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(71) Applicant: **Novartis AG**

4056 Basel (CH)

(72) Inventors:

- **Bagnoli, Fabio**
I-53100 Siena (IT)
- **Biagini, Massimiliano**
I-53100 Siena (IT)

- **Fiaschi, Luigi**
I-53100 Siena (IT)
- **Grandi, Guido**
I-53100 Siena (IT)
- **Mishra, Ravi**
I-53100 Siena (IT)
- **Norais, Nathalie**
I-53100 Siena (IT)
- **Scarselli, Maria**
I-53100 Siena (IT)

(74) Representative: **Carpmaels & Ransford**

**One Southampton Row
London WC1B 5HA (GB)**

Remarks:

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(54) **Compositions for immunising against *Staphylococcus aureus***

(57) An effective *Staphylococcus aureus* vaccine may require several antigenic components, and so various combinations of *S.aureus* antigens are identified for

use in immunisation. These polypeptides may optionally be used in combination with *S.aureus* saccharides.

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Description**TECHNICAL FIELD**

5 **[0001]** This invention relates to antigens derived from *S.aureus* and to their use in immunisation.

BACKGROUND ART

10 **[0002]** *Staphylococcus aureus* is a Gram-positive spherical bacterium. Annual US mortality exceeds that of any other infectious disease, including HIV/AIDS, and *S.aureus* is the leading cause of bloodstream, lower respiratory tract, skin & soft tissue infections. There is currently no authorised vaccine. A vaccine based on a mixture of surface polysaccharides from bacterial types 5 and 8, StaphVAX™, failed to reduce infections when compared to the placebo group in a phase III clinical trial in 2005.

15 **[0003]** Reference 1 reports that the "V710" vaccine from Merck and Intercell is undergoing a phase 2/3 trial on patients undergoing cardiothoracic surgery. The V710 vaccine is based on a single antigen, IsdB [2], a conserved iron-sequestering cell-surface protein.

20 **[0004]** *S.aureus* causes a range of illnesses from minor skin infections to life-threatening diseases such as pneumonia, meningitis, osteomyelitis, bacteremia, endocarditis, toxic shock syndrome, organ abscesses and septicemia. The bacterium has multiple virulence factors which are differentially expressed during different phases of its life cycle, and so a vaccine which can prevent one disease might not prevent another. For instance, the V710 vaccine may be effective against hematic spread of the *S.aureus*, but may be ineffective against pneumonia and may not elicit any opsonic activity. One aim of the invention is to provide vaccines which can protect against hematic spread and pneumonia, and which may also elicit an opsonic response.

25 **[0005]** Thus there remains a need to identify further and improved antigens for use in *S.aureus* vaccines, and in particular for vaccines which are useful against multiple *S.aureus* pathologies.

DISCLOSURE OF THE INVENTION

30 **[0006]** The inventors have identified various *S.aureus* polypeptides that are useful for immunisation, either alone or in combination. These polypeptides may be combined with *S.aureus* saccharides or other *S.aureus* polypeptides. The antigens are useful in *S.aureus* vaccines but may also be used as components in vaccines for immunising against multiple pathogens.

35 **[0007]** The inventors have identified the following 36 polypeptides: clfA, clfB, coA, eap, ebhA, ebpS, efb, emp, esaC, esxA, esxB, FnBA, FnBB, Hla, hlgB, hlgC, isdA, isdB, isdC, isdG, isdH, isdI, lukD, lukE, lukF, lukS, nuc, sasA, sasB, sasC, sasD, sasF, sdrC, sdrD, spa, and sdrE2. This set of antigens is referred to herein as 'the first antigen group'. Thus the invention provides an immunogenic composition comprising a combination of antigens, said combination comprising two or more (i. e. 2, 3, 4, 5, 6 or more) antigens selected from the group consisting of: (1) a clfA antigen; (2) a clfB antigen; (3) a coA antigen; (4) a eap antigen; (5) a ebhA antigen; (6) a ebpS antigen; (7) a efb antigen; (8) a emp antigen; (9) a esaC antigen; (10) a esxA antigen; (11) a esxB antigen; (12) a FnBA antigen; (13) a FnBB antigen; (14) a Hla antigen; (15) a hlgB antigen; (16) a hlgC antigen; (17) a isdA antigen; (18) a isdB antigen; (19) a isdC antigen; (20) a isdG antigen; (21) a isdH antigen; (22) a isdI antigen; (23) a lukD antigen; (24) a lukE antigen; (25) a lukF antigen; (26) a lukS antigen; (27) a nuc antigen; (28) a sasA antigen; (29) a sasB antigen; (30) a sasC antigen; (31) a sasD antigen; (32) a sasF antigen; (33) a sdrC antigen; (34) a sdrD antigen; (35) a spa antigen; (36) a sdrE2 antigen.

45 **[0008]** Within the first antigen group, antigens are preferably selected from a subset of 16 of the 36 polypeptides, namely: clfA, clfB, emp, esaC, esxA, esxB, hla, isdA, isdB, isdC, sasD, sasF, sdrC, sdrD, spa, and sdrE2. Thus the invention provides an immunogenic composition comprising a combination of antigens, said combination comprising two or more (i.e. 2, 3, 4, 5, 6 or more) antigens selected from the group consisting of these sixteen antigens.

50 **[0009]** The inventors have also identified the following 128 polypeptides: sta001, sta002, sta003, sta004, sta005, sta006, sta007, sta008, sta009, sta010, sta011, sta012, sta013, sta014, sta015, sta016, sta017, sta018, sta019, sta020, sta021, sta022, sta023, sta024, sta025, sta026, sta027, sta028, sta029, sta030, sta031, sta032, sta033, sta034, sta035, sta036, sta037, sta038, sta039, sta040, sta041, sta042, sta043, sta044, sta045, sta046, sta047, sta048, sta049, sta050, sta051, sta052, sta053, sta054, sta055, sta056, sta057, sta058, sta059, sta060, sta061, sta062, sta063, sta064, sta065, sta066, sta067, sta068, sta069, sta070, sta071, sta072, sta073, sta074, sta075, sta076, sta077, sta078, sta079, sta080, sta081, sta082, sta083, sta084, sta085, sta086, sta087, sta088, sta089, sta090, sta091, sta092, sta093, sta094, sta095, sta096, sta097, sta098, sta099, sta100, sta101, sta102, sta103, sta104, sta105, sta106, sta107, sta108, sta109, sta110, sta111, sta112, sta113, sta114, sta115, sta116, sta117, sta118, sta119, sta120, NW_6, NW_9, NW_10, NW_7, NW_8, NW_2, NW_1, and NW_5. This set of antigens is referred to herein as 'the second antigen group'. Thus the invention provides an immunogenic composition comprising a combination of antigens, said combination comprising two or more

(i.e. 2, 3, 4, 5, 6 or more) antigens selected from the group consisting of: (1) a sta001 antigen; (2) a sta002 antigen; (3) a sta003 antigen; (4) a sta004 antigen; (5) a sta005 antigen; (6) a sta006 antigen; (7) a sta007 antigen; (8) a sta008 antigen; (9) a sta009 antigen; (10) a sta010 antigen; (11) a sta011 antigen; (12) a sta012 antigen; (13) a sta013 antigen; (14) a sta014 antigen; (15) a sta015 antigen; (16) a sta016 antigen; (17) a sta017 antigen; (18) a sta018 antigen; (19) a sta019 antigen; (20) a sta020 antigen; (21) a sta021 antigen; (22) a sta022 antigen; (23) a sta023 antigen; (24) a sta024 antigen; (25) a sta025 antigen; (26) a sta026 antigen; (27) a sta027 antigen; (28) a sta028 antigen; (29) a sta029 antigen; (30) a sta030 antigen; (31) a sta031 antigen; (32) a sta032 antigen; (33) a sta033 antigen; (34) a sta034 antigen; (35) a sta035 antigen; (36) a sta036 antigen; (37) a sta037 antigen; (38) a sta038 antigen; (39) a sta039 antigen; (40) a sta040 antigen; (41) a sta041 antigen; (42) a sta042 antigen; (43) a sta043 antigen; (44) a sta044 antigen; (45) a sta045 antigen; (46) a sta046 antigen; (47) a sta047 antigen; (48) a sta048 antigen; (49) a sta049 antigen; (50) a sta050 antigen; (51) a sta051 antigen; (52) a sta052 antigen; (53) a sta053 antigen; (54) a sta054 antigen; (55) a sta055 antigen; (56) a sta056 antigen; (57) a sta057 antigen; (58) a sta058 antigen; (59) a sta059 antigen; (60) a sta060 antigen; (61) a sta061 antigen; (62) a sta062 antigen; (63) a sta063 antigen; (64) a sta064 antigen; (65) a sta065 antigen; (66) a sta066 antigen; (67) a sta067 antigen; (68) a sta068 antigen; (69) a sta069 antigen; (70) a sta070 antigen; (71) a sta071 antigen; (72) a sta072 antigen; (73) a sta073 antigen; (74) a sta074 antigen; (75) a sta075 antigen; (76) a sta076 antigen; (77) a sta077 antigen; (78) a sta078 antigen; (79) a sta079 antigen; (80) a sta080 antigen; (81) a sta081 antigen; (82) a sta082 antigen; (83) a sta083 antigen; (84) a sta084 antigen; (85) a sta085 antigen; (86) a sta086 antigen; (87) a sta087 antigen; (88) a sta088 antigen; (89) a sta089 antigen; (90) a sta090 antigen; (91) a sta091 antigen; (92) a sta092 antigen; (93) a sta093 antigen; (94) a sta094 antigen; (95) a sta095 antigen; (96) a sta096 antigen; (97) a sta097 antigen; (98) a sta098 antigen; (99) a sta099 antigen; (100) a sta100 antigen; (101) a sta101 antigen; (102) a sta102 antigen; (103) a sta103 antigen; (104) a sta104 antigen; (105) a sta105 antigen; (106) a sta106 antigen; (107) a sta107 antigen; (108) a sta108 antigen; (109) a sta109 antigen; (110) a sta110 antigen; (111) a sta111 antigen; (112) a sta112 antigen; (113) a sta113 antigen; (114) a sta114 antigen; (115) a sta115 antigen; (116) a sta116 antigen; (117) a sta117 antigen; (118) a sta118 antigen; (119) a sta119 antigen; (120) a sta120 antigen; (121) a NW_6 antigen; (122) a NW_9 antigen; (123) a NW_10 antigen; (124) a NW_7 antigen; (125) a NW_8 antigen; (126) a NW_2 antigen; (127) a NW_1 antigen; and (128) a NW_5 antigen.

[0010] Within the second antigen group of 128 antigens, a preferred subset of 113 antigens omits (81) and (107) to (120) from this list.

[0011] Within the second antigen group, a subset of 27 of the 128 polypeptides is referred to herein as 'the third antigen group', namely: sta001, sta002, sta003, sta004, sta005, sta006, sta007, sta008, sta009, sta010, sta019, sta028, sta040, sta049, sta057, sta064, sta073, sta095, sta098, sta101, sta105, NW_1, NW_6, NW_7, NW_8, NW_9 and NW_10. The invention provides an immunogenic composition comprising a combination of antigens, said combination comprising two or more (i.e. 2, 3, 4, 5, 6 or more) antigens selected from the third antigen group.

[0012] The 101 antigens that are in the second antigen group but not in the third antigen group are referred to herein as 'the fourth antigen group'. Within the fourth antigen group of 101 antigens, a preferred subset of 86 antigens omits (81) and (107) to (120) from the above list. The second antigen group thus consists of a combination of the third and fourth antigen groups.

[0013] Within the second antigen group, a subset of 8 of the 128 polypeptides is referred to herein as 'the fifth antigen group', namely: sta004, sta006, sta007, sta011, sta028, sta060, sta098 and sta112. The invention provides an immunogenic composition comprising a combination of antigens, said combination comprising two or more (i.e. 2, 3, 4, 5, 6 or more) antigens selected from the fifth antigen group.

[0014] Within the 36 antigens of the first antigen group there are 630 possible pairs of different antigens. All such pairs are disclosed herein and are part of the invention. Thus the invention provides an immunogenic composition comprising a pair of antigens, wherein said pair is one of said 630 pairs.

[0015] Within the 128 antigens of the second antigen group there are 8128 possible pairs of different antigens. All such pairs are disclosed herein and are part of the invention. Thus the invention provides an immunogenic composition comprising a pair of antigens, wherein said pair is one of said 8128 pairs.

[0016] Within the preferred 113 antigens of the second antigen group there are 6328 possible pairs of different antigens. All such pairs are disclosed herein and are part of the invention. Thus the invention provides an immunogenic composition comprising a pair of antigens, wherein said pair is one of said 6328 pairs.

[0017] Within the preferred 27 antigens of the third antigen group there are 351 possible pairs of different antigens. All such pairs are disclosed herein and are part of the invention. Thus the invention provides an immunogenic composition comprising a pair of antigens, wherein said pair is one of said 351 pairs.

[0018] Within the 101 antigens of the fourth antigen group there are 5050 possible pairs of different antigens. All such pairs are disclosed herein and are part of the invention. Thus the invention provides an immunogenic composition comprising a pair of antigens, wherein said pair is one of said 5050 pairs.

[0019] Within the preferred 86 antigens of the fourth antigen group there are 3655 possible pairs of different antigens. All such pairs are disclosed herein and are part of the invention. Thus the invention provides an immunogenic composition

comprising a pair of antigens, wherein said pair is one of said 3655 pairs.

[0020] In one embodiment, a composition includes at least one antigen (*i.e.* 1, 2, 3, 4, 5, 6 or more) selected from the first antigen group and at least one antigen (*i.e.* 1, 2, 3, 4, 5, 6 or more) selected from the second antigen group. Antigens from the first antigen group may be selected from the preferred subset of 16 antigens, and antigens from the second antigen group may be selected from the third antigen group or the fifth antigen group.

[0021] The invention also provides an immunogenic composition comprising a combination of antigens, said combination comprising two or more (*i.e.* 2, 3, 4, 5, 6 or more) antigens selected from the group consisting of: (1) a *clfA* antigen; (2) a *clfB* antigen; (3) a *sdrE2* antigen; (4) a *sdrC* antigen; (5) a *SasF* antigen; (6) a *emp* antigen; (7) a *sdrD* antigen; (8) a *spa* antigen; (9) a *esaC* antigen; (10) a *esxA* antigen; (11) a *esxB* antigen; (12) a *sta006* antigen; (13) a *isdC* antigen; (14) a *hla* antigen; (15) a *sta011* antigen; (16) *isdA* antigen; (17) a *isdB* antigen; (18) a *sasF* antigen. This group of 18 antigens is sometimes referred to herein as the 'sixth antigen group'.

[0022] The invention also provides an immunogenic composition comprising a combination of antigens, said combination comprising two or more (*i.e.* 2, 3, 4 or 5) antigens selected from the group consisting of: (1) a *esxA* antigen; (2) a *esxB* antigen; (3) a *sta006* antigen; (4) a *hla* antigen; and/or (5) a *sta011* antigen. The composition may also include an adjuvant *e.g.* an aluminium hydroxide adjuvant.

[0023] Advantageous combinations of the invention are those in which two or more antigens act synergistically. Thus the protection against *S.aureus* disease achieved by their combined administration exceeds that expected by mere addition of their individual protective efficacy.

[0024] Specific combinations of interest include, but are not limited to:

(1) An immunogenic composition comprising a *sdrD* antigen, a *sdrE2* antigen and a *isdC* antigen. The *sdrD* and *sdrE2* antigens can usefully be combined as a hybrid polypeptide, as discussed below, *e.g.* an *SdrDE* hybrid with an *sdrE2* antigen downstream of a *sdrD* antigen.

(2) An immunogenic composition comprising a *sasD* antigen, a *clfB* antigen and a *sdrC* antigen.

(3) An immunogenic composition comprising a *sasD* antigen, a *clfB* antigen, a *sdrC* antigen and a *clfA* antigen.

(4) An immunogenic composition comprising a *sdrD* antigen, a *sdrE2* antigen, a *isdC* antigen and a *sta011* antigen. The *sdrD* and *sdrE2* antigens can usefully be combined as a hybrid polypeptide, as discussed below, *e.g.* a *SdrDE* hybrid with a *sdrE2* antigen downstream of a *sdrD* antigen.

(5) An immunogenic composition comprising a *sasD* antigen, a *clfB* antigen, a *sdrC* antigen and a *sta006* antigen.

(6) An immunogenic composition comprising a *sdrD* antigen, a *sdrE2* antigen, a *isdC* antigen and a *hla* antigen. The *sdrD* and *sdrE2* antigens can usefully be combined as a hybrid polypeptide, as discussed below, *e.g.* a *SdrDE* hybrid with a *sdrE2* antigen downstream of a *sdrD* antigen. The *Hla* antigen may be a detoxified mutant *e.g.* including a H35L mutation.

(7) An immunogenic composition comprising a *sasD* antigen, a *clfB* antigen, a *sdrC* antigen and a *esxA* antigen.

(8) An immunogenic composition comprising a *esxA* antigen, a *esxB* antigen, a *sta006* antigen and a *hla* antigen. The *esxA* and *esxB* antigens can usefully be combined as a hybrid polypeptide, as discussed below, *e.g.* a *EsxAB* hybrid with a *esxB* antigen downstream of a *esxA* antigen. The *Hla* antigen may be a detoxified mutant *e.g.* including a H35L mutation.

(9) An immunogenic composition comprising a *sdrD* antigen, a *sdrE2* antigen, a *isdC* antigen and a *esxA* antigen. The *sdrD* and *sdrE2* antigens can usefully be combined as a hybrid polypeptide, as discussed below, *e.g.* a *SdrDE* hybrid with a *sdrE2* antigen downstream of a *sdrD* antigen.

(10) An immunogenic composition comprising a *esxA* antigen, a *esxB* antigen, a *sta006* antigen and a *sta011* antigen. The *esxA* and *esxB* antigens may be combined as a hybrid polypeptide, as discussed below, *e.g.* an *EsxAB* hybrid.

(11) An immunogenic composition comprising a *esxA* antigen, a *esxB* antigen and a *sta011* antigen. The *esxA* and *esxB* antigens can usefully be combined as a hybrid polypeptide, as discussed below, *e.g.* a *EsxAB* hybrid with a *esxB* antigen downstream of a *esxA* antigen.

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(12) An immunogenic composition comprising a sasD antigen, a clfB antigen, a sdrC antigen and a spa antigen.

5 (13) An immunogenic composition comprising a esxA antigen, a esxB antigen, a isdA antigen, a sta006 antigen, a sta011 antigen and a spa antigen. The esxA and esxB antigens may be combined as a hybrid polypeptide, as discussed below, *e.g.* an EsxAB hybrid. The isdA antigen may be a fragment of a full-length isdA antigen *e.g.* SEQ ID NO: 157. The spa antigen may be a fragment of a full-length spa antigen, such as a Spa(D) domain mutated to disrupt or decrease binding to IgG Fc.

10 (14) An immunogenic composition comprising a esxA antigen, a esxB antigen, a H1a antigen, a sta006 antigen and a sta011 antigen. The esxA and esxB antigens may be combined as a hybrid polypeptide, as discussed below, *e.g.* an EsxAB hybrid. The H1a antigen may be a detoxified mutant *e.g.* including a H35L mutation.

15 (15) An immunogenic composition comprising a sdrD antigen, a sdrE2 antigen, a isdC antigen and a sdrE2 antigen. The sdrD and sdrE2 antigens can usefully be combined as a hybrid polypeptide, as discussed below, *e.g.* a SdrDE hybrid with a sdrE2 antigen downstream of a sdrD antigen.

20 (16) An immunogenic composition comprising a esxA antigen, a esxB antigen and a hla antigen. The esxA and esxB antigens can usefully be combined as a hybrid polypeptide, as discussed below, *e.g.* a EsxAB hybrid with a esxB antigen downstream of a esxA antigen. The H1a antigen may be a detoxified mutant *e.g.* including a H35L mutation.

25 (17) An immunogenic composition comprising a hla antigen, a isdA antigen, a sta006 antigen and a sta011 antigen. The isdA antigen may be a fragment of a full-length isdA antigen *e.g.* SEQ ID NO: 157. The Hla antigen may be a detoxified mutant *e.g.* including a H35L mutation.

30 (18) An immunogenic composition comprising a esxA antigen, a esxB antigen, a sta006 antigen and a isdA antigen. The esxA and esxB antigens can usefully be combined as a hybrid polypeptide, as discussed below, *e.g.* a EsxAB hybrid with a esxB antigen downstream of a esxA antigen. The isdA antigen may be a fragment of a full-length isdA antigen *e.g.* SEQ ID NO: 157.

(19) An immunogenic composition comprising a sasD antigen, a clfB antigen, a sdrC antigen and a hla antigen. The Hla antigen may be a detoxified mutant *e.g.* including a H35L mutation.

35 (20) An immunogenic composition comprising a H1a antigen, a sta006 antigen and a sta011 antigen. The H1a antigen may be a detoxified mutant *e.g.* including a H35L mutation.

40 (21) An immunogenic composition comprising a esxA antigen and a esxB antigen. The esxA and esxB antigens can usefully be combined as a hybrid polypeptide, as discussed below, *e.g.* an EsxAB hybrid with an esxB antigen downstream of an esxA antigen.

(22) An immunogenic composition comprising a esxA antigen, a esxB antigen and a sta006 antigen. The esxA and esxB antigens can usefully be combined as a hybrid polypeptide, as discussed below, *e.g.* a EsxAB hybrid with a esxB antigen downstream of a esxA antigen.

45 (23) An immunogenic composition comprising a spa antigen, a sta006 antigen and a sta011 antigen. The spa antigen may be a fragment of a full-length spa antigen, such as a Spa(D) domain mutated to disrupt or decrease binding to IgG Fc.

50 (24) An immunogenic composition comprising a esxA antigen, a esxB antigen, a isdA antigen, a sta006 antigen and a sta011 antigen. The esxA and esxB antigens may be combined as a hybrid polypeptide, as discussed below, *e.g.* an EsxAB hybrid. The isdA antigen may be a fragment of a full-length isdA antigen *e.g.* SEQ ID NO: 157.

(25) An immunogenic composition comprising a sta006 antigen and a sta011 antigen.

55 (26) An immunogenic composition comprising a esxA antigen, a esxB antigen, a sta006 antigen, a isdA antigen and a clfB antigen. The esxA and esxB antigens can usefully be combined as a hybrid polypeptide, as discussed below, *e.g.* a EsxAB hybrid with a esxB antigen downstream of a esxA antigen. The isdA antigen may be a fragment of a full-length isdA antigen *e.g.* SEQ ID NO: 157. The clfB antigen may be a fragment of a full-length clfB antigen

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e.g. SEQ ID NO: 163.

(27) An immunogenic composition comprising a sta006 antigen, a sta011 antigen and a sta019 antigen.

5 (28) An immunogenic composition comprising a esxA antigen, a esxB antigen, a sta006 antigen, a hla antigen and a clfB antigen. The esxA and esxB antigens can usefully be combined as a hybrid polypeptide, as discussed below, e.g. a EsxAB hybrid with a esxB antigen downstream of a esxA antigen. The clfB antigen may be a fragment of a full-length clfB antigen e.g. SEQ ID NO: 163. The H1a antigen may be a detoxified mutant e.g. including a H35L mutation.

10 (29) An immunogenic composition comprising a sta006 antigen, a sta011 antigen, a sta019 antigen, and a hla antigen. The H1a antigen may be a detoxified mutant e.g. including a H35L mutation.

15 (30) An immunogenic composition comprising a esxA antigen, a esxB antigen, a sta006 antigen, a sta011 antigen and a clfB antigen. The esxA and esxB antigens can usefully be combined as a hybrid polypeptide, as discussed below, e.g. a EsxAB hybrid with a esxB antigen downstream of a esxA antigen. The clfB antigen may be a fragment of a full-length clfB antigen e.g. SEQ ID NO: 163.

20 (31) An immunogenic composition comprising a spa antigen, a esxA antigen, a esxB antigen, a sta006 antigen and a sta011 antigen. The spa antigen may be a fragment of a full-length spa antigen, such as a Spa(D) domain mutated to disrupt or decrease binding to IgG Fc. The esxA and esxB antigens may be combined as a hybrid polypeptide, as discussed below, e.g. an EsxAB hybrid.

25 (32) An immunogenic composition comprising a sdrD antigen, a sdrE2 antigen, a isdC antigen and a esxB antigen. The sdrD and sdrE2 antigens can usefully be combined as a hybrid polypeptide, as discussed below, e.g. a SdrDE hybrid with a sdrE2 antigen downstream of a sdrD antigen.

30 (33) An immunogenic composition comprising a esxA antigen, a esxB antigen, a sta006 antigen, a sta011 antigen and a sta019 antigen. The esxA and esxB antigens can usefully be combined as a hybrid polypeptide, as discussed below, e.g. a EsxAB hybrid with a esxB antigen downstream of a esxA antigen.

35 (34) An immunogenic composition comprising a esxA antigen, a esxB antigen, a sta006 antigen, a isdA antigen and a sdrD antigen. The esxA and esxB antigens can usefully be combined as a hybrid polypeptide, as discussed below, e.g. a EsxAB hybrid with a esxB antigen downstream of a esxA antigen. The isdA antigen may be a fragment of a full-length isdA antigen e.g. SEQ ID NO: 157. The sdrD antigen may be a fragment of a full-length sdrD antigen e.g. SEQ ID NO: 156.

40 (35) An immunogenic composition comprising a esxA antigen, a esxB antigen, and a isdA antigen. The esxA and esxB antigens can usefully be combined as a hybrid polypeptide, as discussed below, e.g. a EsxAB hybrid with a esxB antigen downstream of a esxA antigen. The isdA antigen may be a fragment of a full-length isdA antigen e.g. SEQ ID NO: 157.

45 (36) An immunogenic composition comprising a sasD antigen, a clfB antigen, a sdrC antigen, a esxA antigen and a esxB antigen. The esxA and esxB antigens can usefully be combined as a hybrid polypeptide, as discussed below, e.g. an EsxAB hybrid with an esxB antigen downstream of an esxA antigen.

50 (37) An immunogenic composition comprising a H1a antigen, a spa antigen, a sta006 antigen and a sta011 antigen. The H1a antigen may be a detoxified mutant e.g. including a H35L mutation. The spa antigen may be a fragment of a full-length spa antigen, such as a Spa(D) domain mutated to disrupt or decrease binding to IgG Fc.

[0025] In some embodiments, any of these 37 compositions may include additional staphylococcal antigens, and these further antigens can be polypeptides and/or saccharides. For example, they can usefully also include one or more *S.aureus* capsular saccharide conjugate(s) e.g. against a serotype 5 and/or a serotype 8 strain. The inclusion of one or both such conjugates is particularly useful for combinations (8), (10), (20), (23), (25), (31) and (37).

55 [0026] In other embodiments, these 37 compositions include no additional staphylococcal polypeptide antigens. In other embodiments, these 37 compositions include no additional staphylococcal antigens. In other embodiments, these 37 compositions include no additional antigens.

[0027] The invention also provides a polypeptide comprising amino acid sequence (a) having 80% or more identity

(e.g. 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 151; and/or (b) comprising a fragment of at least 'n' consecutive amino acids from amino acids 1-97 of SEQ ID NO: 151 and at least 'n' consecutive amino acids from amino acids 104-207 of SEQ ID NO: 151, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). The invention also provides a polypeptide comprising amino acid sequence (a) having 80% or more identity (e.g. 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 152; and/or (b) comprising a fragment of at least 'n' consecutive amino acids from amino acids 1-104 of SEQ ID NO: 152 and at least 'n' consecutive amino acids from amino acids 111-207 of SEQ ID NO: 152, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These polypeptides can elicit antibodies (e.g. when administered to a human) which recognise both the wild-type staphylococcal protein comprising SEQ ID NO: 10 and the wild-type staphylococcal protein comprising SEQ ID NO: 11. Thus the immune response will recognise both of antigens *esxA* and *esxB*. Preferred fragments of (b) provide an epitope from SEQ ID NO: 10 and an epitope from SEQ ID NO: 11. The invention also provides an immunogenic composition comprising a combination of such a protein and an adjuvant, such as an aluminium hydroxide adjuvant.

[0028] The invention also provides a polypeptide comprising amino acid sequence (a) having 80% or more identity (e.g. 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 241; and/or (b) comprising both a fragment of at least 'n' consecutive amino acids from amino acids 1-96 of SEQ ID NO: 241 and a fragment of at least 'n' consecutive amino acids from amino acids 103-205 of SEQ ID NO: 241, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These polypeptides (e.g. SEQ ID NO: 250) can elicit antibodies (e.g. when administered to a human) which recognise both the wild-type staphylococcal protein comprising SEQ ID NO: 10 and the wild-type staphylococcal protein comprising SEQ ID NO: 11. Thus the immune response will recognise both of antigens *esxA* and *esxB*. Preferred fragments of (b) provide an epitope from SEQ ID NO: 10 and an epitope from SEQ ID NO: 11. The invention also provides an immunogenic composition comprising a combination of such a protein and an adjuvant, such as an aluminium hydroxide adjuvant.

[0029] The invention also provides a polypeptide comprising a staphylococcal hemolysin sequence, wherein the sequence does not include a sequence having at least 90% identity to SEQ ID NO: 217 but can elicit antibodies which can kill staphylococci. The polypeptide may have a first sequence having 80% or more identity (e.g. 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 218 and a second sequence having 80% or more identity (e.g. 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 219, wherein the first and second sequences are either directly joined or are joined by an intervening amino acid sequence having fewer than 40 amino acids (e.g. ≤ 35 amino acids, ≤ 30 amino acids, ≤ 25 amino acids, ≤ 20 amino acids, ≤ 15 amino acids, ≤ 10 amino acids, ≤ 5 amino acids). SEQ ID NOs: 189 and 216 are examples of such polypeptides, in which the first and second sequences are joined by a tetrapeptide PSGS sequence (SEQ ID NO: 225).

[0030] The invention also provides an immunogenic composition comprising a Sta011 antigen and a Ca^{++} ion. The antigen and Ca^{++} ion may form a complex e.g. atoms in the antigen may coordinate the Ca^{++} ion. The immunogenic composition may also include an adjuvant.

[0031] The invention also provides a oligomer of a Sta011 antigen, and also immunogenic compositions comprising such oligomers. The oligomer can be a dimer, trimer, tetramer, pentamer, hexamer, heptamer, octamer or higher. An oligomer may comprise a Ca^{++} ion, and a composition comprising Sta011 oligomers may comprise 5-500mM Ca^{++} ions.

Further polypeptide antigens

[0032] In additions to antigens from the various antigen groups of the invention, immunogenic compositions may include one or more of the following *S.aureus* antigens (or antigens comprising immunogenic fragment(s) thereof) to enhance the efficacy against *S.aureus* of an immune response elicited by the composition [e.g. see references 3-10]:

- AhpC
- AhpF
- Autolysin amidase
- Autolysin glucosaminidase
- Collagen binding protein CAN
- EbhB

- GehD lipase
- Heparin binding protein HBP (17kDa)
- 5 ● Laminin receptor
- MAP
- 10 ● MntC (also known as SitC)
- MRPII
- Npase
- 15 ● ORF0594
- ORF0657n
- ORF0826
- 20 ● PBP4
- RAP (RNA III activating protein)
- 25 ● Sai-1
- SasK
- SBI
- 30 ● SdrG
- SdrH
- 35 ● SSP-1
- SSP-2
- Vitronectin-binding protein
- 40

Combinations with saccharides

[0033] The individual antigens identified in the antigen groups of the invention may be used in combination with conjugated saccharide antigens. Thus the invention provides an immunogenic composition comprising a combination of:

- (1) one or more antigen(s) selected from the first, second, third or fourth antigen groups (as defined above); and
- (2) one or more conjugates of a *S.aureus* exopolysaccharide and a carrier protein.

[0034] A conjugate used in component (2) of this combination includes a saccharide moiety and a carrier moiety. The saccharide moiety is from the exopolysaccharide of *S.aureus*, which is a poly-N-acetylglucosamine (PNAG). The saccharide may be a polysaccharide having the size that arises during purification of the exopolysaccharide from bacteria, or it may be an oligosaccharide achieved by fragmentation of such a polysaccharide e.g. size can vary from over 400kDa to between 75 and 400kDa, or between 10 and 75kDa, or up to 30 repeat units. The saccharide moiety can have various degrees of N-acetylation and, as described in reference 11, the PNAG may be less than 40% N-acetylated (e.g. less than 35, 30, 20, 15, 10 or 5% N-acetylated; deacetylated PNAG is also known as dPNAG). Deacetylated epitopes of PNAG can elicit antibodies that are capable of mediating opsonic killing. The PNAG may or may not be O-succinylated e.g. it may be O-succinylated on fewer less than 25, 20, 15, 10, 5, 2, 1 or 0.1 % of residues.

[0035] The invention also provides an immunogenic composition comprising a combination of:

- (1) one or more antigen(s) selected from the first, second, third or fourth antigen groups; and
- (2) one or more conjugates of a *S.aureus* capsular saccharide and a carrier protein.

[0036] A conjugate used in component (2) of this combination includes a saccharide moiety and a carrier moiety. The saccharide moiety is from the capsular saccharide of a *S.aureus*. The saccharide may be a polysaccharide having the size that arises during purification of capsular polysaccharide from bacteria, or it may be an oligosaccharide achieved by fragmentation of such a polysaccharide. Capsular saccharides may be obtained from any suitable strain of *S.aureus* (or any bacterium having a similar or identical saccharide), such as from a type 5 and/or a type 8 *S.aureus* strain and/or a type 336 *S.aureus* strain. Most strains of infectious *S.aureus* contain either Type 5 or Type 8 capsular saccharides. Both have FucNAcp in their repeat unit as well as ManNAcA which can be used to introduce a sulfhydryl group for linkage. The repeating unit of the Type 5 saccharide is $\rightarrow 4$)- β -D-Man NAcA-(1 \rightarrow 4)- α -L-FucNAc(30Ac)-(1 \rightarrow 3)- β -D-FucNAc-(1 \rightarrow , whereas the repeating unit of the Type 8 saccharide is $\rightarrow 3$)- β -D-ManNAcA(40Ac)-(1 \rightarrow 3)- α -L-FucNAc (1 \rightarrow 3)- α -D-FucNAc(1 \rightarrow . The type 336 saccharide is a β -linked hexosamine with no O-acetylation [12,13] and is cross-reactive with antibodies raised against the 336 strain (ATCC 55804). A combination of a type 5 and a type 8 saccharide is typical, and a type 336 saccharide may be added to this pairing [14].

[0037] The invention also provides an immunogenic composition comprising a combination of:

- (1) one or more antigen(s) selected from the first, second, third or fourth antigen groups;
- (2) one or more conjugates of a *S.aureus* exopolysaccharide and a carrier protein; and
- (3) one or more conjugates of a *S.aureus* capsular saccharide and a carrier protein.

[0038] The carrier moiety in these conjugates will usually be a protein, but usually not one of the antigens of (1). Typical carrier proteins are bacterial toxins, such as diphtheria or tetanus toxins, or toxoids or mutants or fragments thereof. The CRM197 diphtheria toxin mutant [15] is useful. Other suitable carrier proteins include the *N.meningitidis* outer membrane protein complex [16], synthetic peptides [17,18], heat shock proteins [19,20], pertussis proteins [21,22], cytokines [23], lymphokines [23], hormones [23], growth factors [23], artificial proteins comprising multiple human CD4⁺ T cell epitopes from various pathogen-derived antigens [24] such as N19 [25], protein D from *H.influenzae* [26-28], pneumolysin [29] or its non-toxic derivatives [30], pneumococcal surface protein PspA [31], iron-uptake proteins [32], toxin A or B from *C.difficile* [33], recombinant *P.aeruginosa* exoprotein A (rEPA) [34], etc. In some embodiments the carrier protein is a *S.aureus* protein, such as an antigen selected from the first, second, third or fourth antigen groups.

[0039] Where a composition includes more than one conjugate, each conjugate may use the same carrier protein or a different carrier protein.

[0040] Conjugates may have excess carrier (w/w) or excess saccharide (w/w). In some embodiments, a conjugate may include substantially equal weights of each.

[0041] The carrier molecule may be covalently conjugated to the carrier directly or via a linker. Direct linkages to the protein may be achieved by, for instance, reductive amination between the saccharide and the carrier, as described in, for example, references 35 and 36. The saccharide may first need to be activated e.g. by oxidation. Linkages via a linker group may be made using any known procedure, for example, the procedures described in references 37 and 38. A preferred type of linkage is an adipic acid linker, which may be formed by coupling a free -NH₂ group (e.g. introduced to a glucan by amination) with adipic acid (using, for example, diimide activation), and then coupling a protein to the resulting saccharide-adipic acid intermediate [39,40]. Another preferred type of linkage is a carbonyl linker, which may be formed by reaction of a free hydroxyl group of a saccharide CDI [41, 42] followed by reaction with a protein to form a carbamate linkage. Other linkers include β -propionamido [43], nitrophenyl-ethylamine [44], haloacyl halides [45], glycosidic linkages [46], 6-aminocaproic acid [47], ADH [48], C₄ to C₁₂ moieties [49], etc. Carbodiimide condensation can also be used [50].

[0042] PNAG conjugates may be prepared in various ways e.g. by a process comprising: a) activating the PNAG by adding a linker comprising a maleimide group to form an activated PNAG; b) activating the carrier protein by adding a linker comprising a sulphhydryl group to form an activated carrier protein; and c) reacting the activated PNAG and the activated carrier protein to form a PNAG-carrier protein conjugate; or by a process comprising a) activating the PNAG by adding a linker comprising a sulphhydryl group to form an activated PNAG; b) activating the carrier protein by adding a linker comprising a maleimide group to form an activated carrier protein; and c) reacting the activated PNAG and the activated carrier protein to form a PNAG-carrier protein conjugate; or by a process comprising a) activating the PNAG by adding a linker comprising a sulphhydryl group to form an activated PNAG; b) activating the carrier protein by adding a linker comprising a sulphhydryl group to form an activated carrier protein; and c) reacting the activated PNAG and the activated carrier protein to form a PNAG-carrier protein conjugate.

[0043] The individual antigens identified in the antigen groups of the invention may be used as carrier proteins for

exopolysaccharides, to form a covalent conjugate. Thus the invention provides an immunogenic composition comprising a conjugate of (1) an antigen selected from the first, second, third and fourth antigen groups and (2) a *S.aureus* exopolysaccharide. The invention also provides an immunogenic composition comprising a conjugate of (1) an antigen selected from the first, second, third and fourth antigen groups and (2) a *S.aureus* capsular saccharide. Further characteristics of such conjugates are described above. These conjugates may be combined with any of the antigens disclosed herein.

Combinations with non-staphylococcal antigens

[0044] The individual antigens identified in the antigen groups of the invention may be used in combination with non-staphylococcal antigens, and in particular with antigens from bacteria associated with nosocomial infections. Thus the invention provides an immunogenic composition comprising a combination of:

- (1) one or more antigen(s) selected from the first, second, third and fourth antigen groups (as defined above); and
- (2) one or more antigen(s) selected from the group consisting of: *Clostridium difficile*; *Pseudomonas aeruginosa*; *Candida albicans*; and extraintestinal pathogenic *Escherichia coli*.

[0045] Further suitable antigens for use in combination with staphylococcal antigens of the invention are listed on pages 33-46 of reference 51.

First antigen group

clfA

[0046] The '*clfA*' antigen is annotated as 'clumping factor A'. In the NCTC 8325 strain *clfA* is SAOUHSC_00812 and has amino acid sequence SEQ ID NO: 1 (GI:88194572). In the Newman strain it is *nwmn_0756* (GI:151220968).

[0047] Useful *clfA* antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 1 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 1; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 1, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These *clfA* proteins include variants of SEQ ID NO: 1. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 1. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 1 while retaining at least one epitope of SEQ ID NO: 1. The final 368 C-terminal amino acids of SEQ ID NO: 1 can usefully be omitted. The first 39 N-terminal amino acids of SEQ ID NO: 1 can usefully be omitted. Other fragments omit one or more protein domains.

[0048] SEQ ID NO: 224 is a useful fragment of SEQ ID NO: 1 ('Clf₄₀₋₅₅₉'). This fragment omits the long repetitive region towards the C-terminal of SEQ ID NO: 1.

clfB

[0049] The '*clfB*' antigen is annotated as 'clumping factor B'. In the NCTC 8325 strain *clfB* is SAOUHSC_02963 and has amino acid sequence SEQ ID NO: 2 (GI:88196585). In the Newman strain it is *nwmn_2529* (GI:151222741).

[0050] Useful *clfB* antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 2 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 2; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 2, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These *clfB* proteins include variants of SEQ ID NO: 2. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 2. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 2 while retaining at least one epitope of SEQ ID NO: 2. The final 40 C-terminal amino acids of SEQ ID NO: 2 can usefully be omitted. The first 44 N-terminal amino acids of SEQ ID NO: 2 can usefully be omitted. Other fragments omit one or more protein domains. ClfB is naturally a long protein and so the use of fragments is helpful e.g. for purification, handling, fusion, expression, etc. SEQ ID NO: 163 is a useful fragment of SEQ ID NO: 2 ('ClfB₄₅₋₅₅₂'). This fragment includes the most exposed domain of ClfB and is more easily used at an industrial scale. It also reduces the antigen's similarity with human proteins. Other useful fragments, based on a 3-domain model of ClfB, include: ClfB₄₅₋₃₆₀ (also known as CLFB-N12; SEQ ID NO: 196); ClfB₂₁₂₋₅₄₂ (also known as CLFB-N23; SEQ ID NO: 197); and ClfB₃₆₀₋₅₄₂ (also known as CLFB-N3; SEQ ID NO: 198).

coA

[0051] The 'coA' antigen is annotated as 'coagulase Coa'. In the NCTC 8325 strain coA is SAOUHSC_00192 and has amino acid sequence SEQ ID NO: 3 (GI:88194002). In the Newman strain it is nwmn_0166 (GI:151220378).

[0052] Useful coA antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 3 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 3; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 3, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These coA proteins include variants of SEQ ID NO: 3. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 3. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 3 while retaining at least one epitope of SEQ ID NO: 3. The first 14 N-terminal amino acids of SEQ ID NO: 3 can usefully be omitted. Other fragments omit one or more protein domains.

eap

[0053] The 'eap' antigen is annotated as 'MHC class II analog protein'. In the NCTC 8325 strain eap is SAOUHSC_02161 and has amino acid sequence SEQ ID NO: 4 (GI:88195840). In the Newman strain it is nwmn_1872 (GI:151222084).

[0054] Useful eap antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 4 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 4; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 4, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These eap proteins include variants of SEQ ID NO: 4. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 4. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 4 while retaining at least one epitope of SEQ ID NO: 4. The first 17 N-terminal amino acids of SEQ ID NO: 4 can usefully be omitted. Other fragments omit one or more protein domains.

ebhA

[0055] The 'ebhA' antigen is annotated as 'EbhA'. In the NCTC 8325 strain ebhA is SAOUHSC_01447 and has amino acid sequence SEQ ID NO: 5 (GI:88195168).

[0056] Useful ebhA antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 5 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 5; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 5, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These ebhA proteins include variants of SEQ ID NO: 5. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 5. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 5 while retaining at least one epitope of SEQ ID NO: 5. The first 39 N-terminal amino acids of SEQ ID NO: 5 can usefully be omitted. Other fragments omit one or more protein domains.

ebpS

[0057] The 'ebpS' antigen is annotated as 'elastin binding protein EbpS'. In the NCTC 8325 strain ebpS is SAOUHSC_01501 and has amino acid sequence SEQ ID NO: 6 (GI:88195217). In the Newman strain it is nwmn_1389 (GI:151221601).

[0058] Useful ebpS antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 6 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 6; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 6, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These ebpS proteins include variants of SEQ ID NO: 6. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 6. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids

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(e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 6 while retaining at least one epitope of SEQ ID NO: 6. Other fragments omit one or more protein domains.

[0059] SEQ ID NO: 165 is a useful fragment of SEQ ID NO: 6 ('EbpS₁₋₁₉₈'). This fragment includes the most exposed domain of EbpS and is more easily used at an industrial scale. It also reduces the antigen's similarity with human proteins.

efb

[0060] The 'efb' antigen is annotated as 'fibrinogen-binding protein truncated'. In the NCTC 8325 strain *efb* is SAOUHSC_01114 and has amino acid sequence SEQ ID NO: 7 (GI:88194860). In the Newman strain it is *nwmn_1069* (GI:151221281).

[0061] Useful *efb* antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 7 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 7; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 7, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150 or more). These *efb* proteins include variants of SEQ ID NO: 7. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 7. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 7 while retaining at least one epitope of SEQ ID NO: 7. The first 14 N-terminal amino acids of SEQ ID NO: 7 can usefully be omitted. Other fragments omit one or more protein domains.

emp

[0062] The 'emp' antigen is annotated as 'extracellular matrix and plasma binding protein'. In the NCTC 8325 strain *emp* is SAOUHSC_00816 and has amino acid sequence SEQ ID NO: 8 (GI:88194575). In the Newman strain it is *nwmn_0758* (GI:151220970).

[0063] Useful *emp* antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 8 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 8; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 8, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These *emp* proteins include variants of SEQ ID NO: 8. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 8. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 8 while retaining at least one epitope of SEQ ID NO: 8. The first 26 N-terminal amino acids of SEQ ID NO: 8 can usefully be omitted. Other fragments omit one or more protein domains.

[0064] SEQ ID NOs: 190, 191, 192 and 193 are useful fragments of SEQ ID NO: 8 ('Emp₃₅₋₃₄₀', 'Emp₂₇₋₃₃₄', 'EMP₃₅₋₃₃₄' and 'Emp₂₇₋₁₄₇', respectively).

esaC

[0065] The 'esaC' antigen is annotated as 'esaC'. In the NCTC 8325 strain *esaC* is SAOUHSC_00264 and has amino acid sequence SEQ ID NO: 9 (GI:88194069).

[0066] Useful *esaC* antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 9 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 9; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 9, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100 or more). These *esaC* proteins include variants of SEQ ID NO: 9. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 9. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 9 while retaining at least one epitope of SEQ ID NO: 9. Other fragments omit one or more protein domains.

esxA

[0067] The 'esxA' antigen is annotated as 'protein'. In the NCTC 8325 strain *esxA* is SAOUHSC_00257 and has amino acid sequence SEQ ID NO: 10 (GI:88194063).

[0068] Useful *esxA* antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO:

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10 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 10; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 10, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90 or more). These esxA proteins include variants of SEQ ID NO: 10. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 10. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 10 while retaining at least one epitope of SEQ ID NO: 10. Other fragments omit one or more protein domains.

5 esxB

[0069] The 'esxB' antigen is annotated as 'esxB'. In the NCTC 8325 strain esxB is SAOUHSC_00265 and has amino acid sequence SEQ ID NO: 11 (GI:88194070).

[0070] Useful esxB antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 11 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 11; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 11, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100 or more). These esxB proteins include variants of SEQ ID NO: 11. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 11. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 11 while retaining at least one epitope of SEQ ID NO: 11. Other fragments omit one or more protein domains.

25 FnBA

[0071] The 'FnBA' antigen is annotated as 'fibronectin-binding protein A precursor FnBPA'. In the NCTC 8325 strain FnBA is SAOUHSC_02803 and has amino acid sequence SEQ ID NO: 12 (GI:88196438). In the Newman strain it is nwmn_2399 (GI:151222611). Proteomic analysis has revealed that this protein is secreted or surface-exposed.

[0072] Useful FnBA antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 12 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 12; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 12, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These FnBA proteins include variants of SEQ ID NO: 12. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 12. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 12 while retaining at least one epitope of SEQ ID NO: 12. The final 37 C-terminal amino acids of SEQ ID NO: 12 can usefully be omitted. Other fragments omit one or more protein domains. FnBA is naturally a long protein and so the use of fragments is helpful e.g. for purification, handling, fusion, expression, etc.

[0073] SEQ ID NOs: 166 ('FnBA₁₋₅₁₁') and 167 ('FnBA₅₁₂₋₉₅₃') are useful fragments of SEQ ID NO: 12. These fragments are more easily used at an industrial scale.

FnBB

[0074] The 'FnBB' antigen is annotated as 'fibronectin binding protein B FnBPB'. In the NCTC 8325 strain FnBB is SAOUHSC_02802 and has amino acid sequence SEQ ID NO: 13 (GI:88196437). In the Newman strain it is nwmn_2397 (GI:151222609).

[0075] Useful FnBB antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 13 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 13; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 13, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These FnBB proteins include variants of SEQ ID NO: 13. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 13. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 13 while retaining at least one epitope of SEQ ID NO: 13. The final 37 C-terminal amino acids of SEQ ID NO: 13 can usefully be omitted. Other fragments omit one or more protein domains.

Hla

[0076] The 'Hla' antigen is the 'alpha-hemolysin precursor' also known as 'alpha toxin' or simply 'hemolysin'. In the NCTC 8325 strain Hla is SAOUHSC_01121 and has amino acid sequence SEQ ID NO: 14 (GI:88194865). In the Newman strain it is nwmn_1073 (GI:151221285). Hla is an important virulence determinant produced by most strains of *S. aureus*, having pore-forming and haemolytic activity. Anti-Hla antibodies can neutralise the detrimental effects of the toxin in animal models, and Hla is particularly useful for protecting against pneumonia.

[0077] Useful Hla antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 14 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 14; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 14, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These Hla proteins include variants of SEQ ID NO: 14. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 14. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 14 while retaining at least one epitope of SEQ ID NO: 14. The first 26 N-terminal amino acids of SEQ ID NO: 14 can usefully be omitted (e.g. to give SEQ ID NO: 231). Truncation at the C-terminus can also be used e.g. leaving only 50 amino acids (residues 27-76 of SEQ ID NO: 14) [52]. Other fragments omit one or more protein domains.

[0078] Hla's toxicity can be avoided in compositions of the invention by chemical inactivation (e.g. using formaldehyde, glutaraldehyde or other cross-linking reagents). Instead, however, it is preferred to use mutant forms of Hla which remove its toxic activity while retaining its immunogenicity. Such detoxified mutants are already known in the art. One useful Hla antigen has a mutation at residue 61 of SEQ ID NO: 14, which is residue 35 of the mature antigen (*i.e.* after omitting the first 26 N-terminal amino acids = residue 35 of SEQ ID NO: 231). Thus residue 61 may not be histidine, and may instead be e.g. Ile, Val or preferably Leu. A His-Arg mutation at this position can also be used. For example, SEQ ID NO: 150 is the mature mutant Hla-H35L sequence (*i.e.* SEQ ID NO: 231 with a H35L mutation) and a useful Hla antigen comprises SEQ ID NO: 150. Another useful mutation replaces a long loop with a short sequence e.g. to replace the 39mer at residues 136-174 of SEQ ID NO: 14 with a tetramer such as PSGS (SEQ ID NO: 225), as in SEQ ID NO: 189 (which also includes the H35L mutation) and SEQ ID NO: 216 (which does not include the H35L mutation). Another useful mutation replaces residue Y101 e.g. with a leucine (SEQ ID NO: 242). Another useful mutation replaces residue D152 e.g. with a leucine (SEQ ID NO: 243). Another useful mutant replaces residues H35 and Y101 e.g. with a leucine (SEQ ID NO: 244). Another useful mutant replaces residues H35 and D152 e.g. with a leucine (SEQ ID NO: 245).

[0079] Further useful Hla antigens are disclosed in references 53 and 54.

[0080] SEQ ID NOs: 160, 161 & 194 are three useful fragments of SEQ ID NO: 14 ('Hla₂₇₋₇₆', 'Hla₂₇₋₈₉' and 'Hla₂₇₋₇₉', respectively). SEQ ID NOs: 158, 159 and 195 are the corresponding fragments from SEQ ID NO: 150.

[0081] One useful Hla sequence is SEQ ID NO: 232, which was used in the examples. It has a N-terminal Met, then an Ala-Ser dipeptide from the expression vector, then SEQ ID NO: 150 (from NCTC8325 strain). It is encoded by SEQ ID NO: 233.

hlgB

[0082] The 'hlgB' antigen is annotated as 'leukocidin f subunit precursor HlgB'. In the NCTC 8325 strain hlgB is SAOUHSC_02710 and has amino acid sequence SEQ ID NO: 15 (GI:88196350).

[0083] Useful hlgB antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 15 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 15; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 15, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These hlgB proteins include variants of SEQ ID NO: 15. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 15. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 15 while retaining at least one epitope of SEQ ID NO: 15. The first 26 N-terminal amino acids of SEQ ID NO: 15 can usefully be omitted. Other fragments omit one or more protein domains.

hlgC

[0084] The 'hlgC' antigen is annotated as 'leukocidin s subunit precursor HlgC'. In the NCTC 8325 strain hlgC is SAOUHSC_02709 and has amino acid sequence SEQ ID NO: 16 (GI:88196349).

[0085] Useful hlgC antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO:

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16 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 16; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 16, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These hlgC proteins include variants of SEQ ID NO: 16. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 16. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 16 while retaining at least one epitope of SEQ ID NO: 16. The first 29 N-terminal amino acids of SEQ ID NO: 16 can usefully be omitted. Other fragments omit one or more protein domains.

isdA

[0086] The 'isdA' antigen is annotated as 'IsdA protein'. In the NCTC 8325 strain *isdA* is SAOUHSC_01081 and has amino acid sequence SEQ ID NO: 17 (GI:88194829). In the Newman strain it is *nwmn_1041* (GI: 151221253).

[0087] Useful *isdA* antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 17 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 17; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 17, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These *isdA* proteins include variants of SEQ ID NO: 17. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 17. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 17 while retaining at least one epitope of SEQ ID NO: 17. The final 38 C-terminal amino acids of SEQ ID NO: 17 can usefully be omitted. The first 46 N-terminal amino acids of SEQ ID NO: 17 can usefully be omitted. Truncation to exclude the C-terminal 38mer of SEQ ID NO: 17 (beginning with the LPKTG motif) is also useful. Other fragments omit one or more protein domains.

[0088] SEQ ID NO: 157 is a useful fragment of SEQ ID NO: 17 (amino acids 40-184 of SEQ ID NO: 17; 'IsdA₄₀₋₁₈₄') which includes the natural protein's heme binding site and includes the antigen's most exposed domain. It also reduces the antigen's similarity with human proteins. Other useful fragments are disclosed in references 55 and 56.

[0089] *IsdA* does not adsorb well to aluminium hydroxide adjuvants, so *IsdA* present in a composition may be unadsorbed or may be adsorbed to an alternative adjuvant e.g. to an aluminium phosphate.

[0090] Anti-*IsdA* antibodies protect mice against *S.aureus* abscess formation and lethal challenge [57].

isdB

[0091] The 'isdB' antigen is annotated as 'neurofilament protein *isdB*'. In the NCTC 8325 strain *isdB* is SAOUHSC_01079 and has amino acid sequence SEQ ID NO: 18 (GI:88194828). *IsdB* has been proposed for use as a vaccine antigen on its own [2], but this may not prevent pneumonia.

[0092] Useful *isdB* antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 18 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 18; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 18, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These *isdB* proteins include variants of SEQ ID NO: 18. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 18. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 18 while retaining at least one epitope of SEQ ID NO: 18. The final 36 C-terminal amino acids of SEQ ID NO: 18 can usefully be omitted. The first 40 N-terminal amino acids of SEQ ID NO: 18 can usefully be omitted. Other fragments omit one or more protein domains. Useful fragments of *IsdB* are disclosed in references 56 and 58 e.g. lacking 37 internal amino acids of SEQ ID NO: 18.

[0093] Anti-*IsdB* antibodies protect mice against *S.aureus* abscess formation and lethal challenge [57].

[0094] In some embodiments, compositions of the invention do not include an *isdB* antigen.

isdC

[0095] The 'isdC' antigen is annotated as 'protein'. In the NCTC 8325 strain *isdC* is SAOUHSC_01082 and has amino acid sequence SEQ ID NO: 19 (GI:88194830).

[0096] Useful *isdC* antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 19 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 19; and/or (b) comprising

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a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 19, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These *isdC* proteins include variants of SEQ ID NO: 19. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 19. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 19 while retaining at least one epitope of SEQ ID NO: 19. The final 39 C-terminal amino acids of SEQ ID NO: 19 can usefully be omitted. The first 28 N-terminal amino acids of SEQ ID NO: 19 can usefully be omitted. Other fragments omit one or more protein domains. Useful fragments of *IsdB* are disclosed in reference 56.

[0097] Reference 59 discloses antigens which usefully include epitopes from both *IsdB* and *IsdH*.

isdG

[0098] The '*isdG*' antigen is annotated as 'heme-degrading monooxygenase *IsdG*'. In the NCTC 8325 strain *isdG* is SAOUHSC_01089 and has amino acid sequence SEQ ID NO: 20 (GI:88194836).

[0099] Useful *isdG* antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 20 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 20; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 20, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100 or more). These *isdG* proteins include variants of SEQ ID NO: 20. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 20. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 20 while retaining at least one epitope of SEQ ID NO: 20. Other fragments omit one or more protein domains.

isdH

[0100] The '*isdH*' antigen is annotated as '*isdH*'. In the NCTC 8325 strain *isdH* is SAOUHSC_01843 and has amino acid sequence SEQ ID NO: 21 (GI:88195542). In the Newman strain it is *nwmn_1624* (GI:151221836). It has also been known as *HarA*.

[0101] Useful *isdH* antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 21 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 21; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 21, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These *isdH* proteins include variants of SEQ ID NO: 21. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 21. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 21 while retaining at least one epitope of SEQ ID NO: 21. The final 35 C-terminal amino acids of SEQ ID NO: 21 can usefully be omitted. The first 40 N-terminal amino acids of SEQ ID NO: 21 can usefully be omitted. Other fragments omit one or more protein domains.

[0102] Reference 59 discloses antigens which usefully include epitopes from both *IsdB* and *IsdH*.

isdI

[0103] The '*isdI*' antigen is annotated as 'heme-degrading monooxygenase *IsdI*'. In the NCTC 8325 strain *isdI* is SAOUHSC_00130 and has amino acid sequence SEQ ID NO: 22 (GI:88193943).

[0104] Useful *isdI* antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 22 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 22; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 22, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100 or more). These *isdI* proteins include variants of SEQ ID NO: 22. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 22. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 22 while retaining at least one epitope of SEQ ID NO: 22. Other fragments omit one or more protein domains.

lukD

[0105] The '*lukD*' antigen is annotated as 'leukotoxin *LukD*'. In the NCTC 8325 strain *lukD* is SAOUHSC_01954 and

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has amino acid sequence SEQ ID NO: 23 (GI:88195647). In the Newman strain it is nwmn_1718 (GI:151221930).

[0106] Useful lukD antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 23 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 23; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 23, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These lukD proteins include variants of SEQ ID NO: 23. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 23. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 23 while retaining at least one epitope of SEQ ID NO: 23. The final 43 C-terminal amino acids of SEQ ID NO: 23 can usefully be omitted. The first 26 N-terminal amino acids of SEQ ID NO: 23 can usefully be omitted. Other fragments omit one or more protein domains.

lukE

[0107] The 'lukE' antigen is annotated as 'leukotoxin LukE'. In the NCTC 8325 strain lukE is SAOUHSC_01955 and has amino acid sequence SEQ ID NO: 24 (GI:88195648).

[0108] Useful lukE antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 24 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 24; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 24, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These lukE proteins include variants of SEQ ID NO: 24. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 24. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 24 while retaining at least one epitope of SEQ ID NO: 24. Other fragments omit one or more protein domains.

lukF

[0109] The 'lukF' antigen is annotated as 'Leukocidin/Hemolysin toxin family LukF'. In the NCTC 8325 strain lukF is SAOUHSC_02241 and has amino acid sequence SEQ ID NO: 25 (GI:88195914). Useful lukF antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 25 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 25; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 25, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These lukF proteins include variants of SEQ ID NO: 25. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 25. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 25 while retaining at least one epitope of SEQ ID NO: 25. Other fragments omit one or more protein domains.

lukS

[0110] The 'lukS' antigen is annotated as 'probable leukocidin S subunit LukS'. In the NCTC 8325 strain lukS is SAOUHSC_02243 and has amino acid sequence SEQ ID NO: 26 (GI:88195915). In the Newman strain it is nwmn_1928 (GI: 151222140).

[0111] Useful lukS antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 26 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 26; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 26, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These lukS proteins include variants of SEQ ID NO: 26. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 26. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 26 while retaining at least one epitope of SEQ ID NO: 26. The first 22 N-terminal amino acids of SEQ ID NO: 26 can usefully be omitted. Other fragments omit one or more protein domains.

nuc

[0112] The 'nuc' antigen is annotated as 'thermonuclease precursor'. In the NCTC 8325 strain nuc is SAOUHSC_01316 and has amino acid sequence SEQ ID NO: 27 (GI:88195046).

[0113] Useful nuc antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 27 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 27; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 27, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150 or more). These nuc proteins include variants of SEQ ID NO: 27. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 27. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 27 while retaining at least one epitope of SEQ ID NO: 27. The final 39 C-terminal amino acids of SEQ ID NO: 27 can usefully be omitted. The first 19 N-terminal amino acids of SEQ ID NO: 27 can usefully be omitted. Other fragments omit one or more protein domains.

sasA

[0114] The 'sasA' antigen is annotated as 'SasA'. In the NCTC 8325 strain sasA is SAOUHSC_02990 and has amino acid sequence SEQ ID NO: 28 (GI:88196609).

[0115] Useful sasA antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 28 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 28; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 28, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sasA proteins include variants of SEQ ID NO: 28. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 28. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 28 while retaining at least one epitope of SEQ ID NO: 28. The final 43 C-terminal amino acids of SEQ ID NO: 28 can usefully be omitted. The first 90 N-terminal amino acids of SEQ ID NO: 28 can usefully be omitted. Other fragments omit one or more protein domains.

sasB

[0116] The 'sasB' antigen is annotated as 'fmtB protein; SasB'. In the NCTC 8325 strain sasB is SAOUHSC_02404 and has amino acid sequence SEQ ID NO: 29 (GI:88196065).

[0117] Useful sasB antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 29 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 29; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 29, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sasB proteins include variants of SEQ ID NO: 29. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 29. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 29 while retaining at least one epitope of SEQ ID NO: 29. The final 39 C-terminal amino acids of SEQ ID NO: 29 can usefully be omitted. The first 38 N-terminal amino acids of SEQ ID NO: 29 can usefully be omitted. Other fragments omit one or more protein domains.

sasC

[0118] The 'sasC' antigen is annotated as 'Mrp protein; SasC'. In the NCTC 8325 strain sasC is SAOUHSC_01873 and has amino acid sequence SEQ ID NO: 30 (GI:88195570).

[0119] Useful sasC antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 30 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 30; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 30, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sasC proteins include variants of SEQ ID NO: 30. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 30. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 30 while retaining at least one epitope of SEQ ID NO: 30. The final 36 C-terminal amino acids of SEQ ID NO: 30 can usefully be omitted. The first 37

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N-terminal amino acids of SEQ ID NO: 30 can usefully be omitted. Other fragments omit one or more protein domains.

sasD

5 **[0120]** The 'sasD' antigen is annotated as 'SasD protein'. In the NCTC 8325 strain sasD is SAOUHSC_00094 and has amino acid sequence SEQ ID NO: 31 (GI:88193909).

[0121] Useful sasD antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 31 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 31; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 31, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150 or more). These sasD proteins include variants of SEQ ID NO: 31. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 31. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 31 while retaining at least one epitope of SEQ ID NO: 31. The first 28 N-terminal amino acids of SEQ ID NO: 31 can usefully be omitted. Other fragments omit one or more protein domains.

sasF

20 **[0122]** The 'sasF' antigen is annotated as 'sasF protein'. In the NCTC 8325 strain sasF is SAOUHSC_02982 and has amino acid sequence SEQ ID NO: 32 (GI:88196601).

[0123] Useful sasF antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 32 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 32; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 32, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sasF proteins include variants of SEQ ID NO: 32. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 32. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 32 while retaining at least one epitope of SEQ ID NO: 32. The final 39 C-terminal amino acids of SEQ ID NO: 32 can usefully be omitted. The first 37 N-terminal amino acids of SEQ ID NO: 32 can usefully be omitted. Other fragments omit one or more protein domains.

sdrC

35 **[0124]** The 'sdrC' antigen is annotated as 'sdrC protein'. In the NCTC 8325 strain sdrC is SAOUHSC_00544 and has amino acid sequence SEQ ID NO: 33 (GI:88194324).

[0125] Useful sdrC antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 33 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 33; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 33, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sdrC proteins include variants of SEQ ID NO: 33. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 33. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 33 while retaining at least one epitope of SEQ ID NO: 33. The final 38 C-terminal amino acids of SEQ ID NO: 33 can usefully be omitted. The first 50 N-terminal amino acids of SEQ ID NO: 33 can usefully be omitted. Other fragments omit one or more protein domains. SdrC is naturally a long protein and so the use of fragments is helpful e.g. for purification, handling, fusion, expression, etc.

[0126] SEQ ID NO: 164 is a useful fragment of SEQ ID NO: 33 ('SdrC₅₁₋₅₁₈'). This fragment includes the most exposed domain of SdrC and is more easily used at an industrial scale. It also reduces the antigen's similarity with human proteins.

sdrD

[0127] The 'sdrD' antigen is annotated as 'sdrD protein'. In the NCTC 8325 strain sdrD is SAOUHSC_00545 and has amino acid sequence SEQ ID NO: 34 (GI:88194325).

55 **[0128]** Useful sdrD antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 34 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 34; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 34, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18,

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20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sdrD proteins include variants of SEQ ID NO: 34. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 34. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 34 while retaining at least one epitope of SEQ ID NO: 34. The final 38 C-terminal amino acids of SEQ ID NO: 34 can usefully be omitted. The first 52 N-terminal amino acids of SEQ ID NO: 34 can usefully be omitted. Other fragments omit one or more protein domains. SdrD is naturally a long protein and so the use of fragments is very helpful e.g. for purification, handling, fusion, expression, etc.

[0129] SEQ ID NO: 156 is a useful fragment of SEQ ID NO: 34 ('SdrD₅₃₋₅₉₂'). This fragment includes the most exposed domain of SdrD and is more easily used at an industrial scale. It also reduces the antigen's similarity with human proteins. Another useful fragment, with the same C-terminus residue, is SdrD₃₉₄₋₅₉₂ (also known as SdrD-N3; SEQ ID NO: 199). Another useful fragment is SEQ ID NO: 236 (amino acids 593-1123 of SEQ ID NO: 34), referred to herein as 'SdrD_{CnaB}'.

sdrE2

[0130] The 'sdrE2' antigen is annotated as 'Ser-Asp rich fibrinogen/bone sialoprotein-binding protein SdrE'. In the Newman strain sdrE2 is NWMN_0525 and has amino acid sequence SEQ ID NO: 35 (GI: 151220737).

[0131] Useful sdrE2 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 35 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 35; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 35, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sdrE2 proteins include variants of SEQ ID NO: 35. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 35. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 35 while retaining at least one epitope of SEQ ID NO: 35. The final 38 C-terminal amino acids of SEQ ID NO: 35 can usefully be omitted. The first 52 N-terminal amino acids of SEQ ID NO: 35 can usefully be omitted. Other fragments omit one or more protein domains. SdrE2 is naturally a long protein and so the use of fragments is very helpful e.g. for purification, handling, fusion, expression, etc.

[0132] SEQ ID NO: 155 is a useful fragment of SEQ ID NO: 35 ('SdrE₅₃₋₆₃₂'). This fragment includes the most exposed domain of SdrE2 and is more easily used at an industrial scale. It also reduces the antigen's similarity with human proteins.

spa

[0133] The 'spa' antigen is annotated as 'protein A' or 'SpA'. In the NCTC 8325 strain spa is SAOUHSC_00069 and has amino acid sequence SEQ ID NO: 36 (GI:88193885). In the Newman strain it is nwmn_0055 (GI:151220267). All *S.aureus* strains express the structural gene for spa, a well characterized virulence factor whose cell wall-anchored surface protein product has five highly homologous immunoglobulin binding domains designated E, D, A, B, and C [60]. These domains display ~80% identity at the amino acid level, are 56 to 61 residues in length, and are organized as tandem repeats [61]. SpA is synthesized as a precursor protein with an N-terminal signal peptide and a C-terminal sorting signal [62,63]. Cell wall-anchored spa is displayed in great abundance on the staphylococcal surface [64,65]. Each of its immunoglobulin binding domains is composed of antiparallel α -helices that assemble into a three helix bundle and can bind the Fc domain of immunoglobulin G (IgG) [66,67], the VH3 heavy chain (Fab) of IgM (i.e. the B cell receptor) [68], the von Willebrand factor at its A1 domain [69] and/or the TNF- α receptor I (TNFRI) [70], which is displayed on surfaces of airway epithelia.

[0134] Useful spa antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 36 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 36; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 36, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These spa proteins include variants of SEQ ID NO: 36. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 36. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 36 while retaining at least one epitope of SEQ ID NO: 36. The final 35 C-terminal amino acids of SEQ ID NO: 36 can usefully be omitted. The first 36 N-terminal amino acids of SEQ ID NO: 36 can usefully be omitted. Other fragments omit one or more protein domains. Reference 71 suggests that individual IgG-binding domains might be useful immunogens, alone or in combination.

[0135] SEQ ID NO: 162 is a useful fragment of SEQ ID NO: 36 ('Spa₃₇₋₃₂₅'). This fragment contains all the five SpA Ig-binding domains (which are naturally arranged from N- to C-terminus in the order E, D, A, B, C) and includes the

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most exposed domain of SpA. It also reduces the antigen's similarity with human proteins. Other useful fragments may omit 1, 2, 3 or 4 of the natural A, B, C, D and/or E domains to prevent the excessive B cell expansion and then apoptosis which might occur if spa functions as a B cell superantigen. As reported in reference 71, other useful fragments may include only 1, 2, 3 or 4 of the natural A, B, C, D and/or E domains *e.g.* comprise only the SpA(A) domain but not B to E, or comprise only the SpA(D) domain but not A, B, C or E, *etc.* Thus a spa antigen useful with the invention may include 1, 2, 3, 4 or 5 IgG-binding domains, but ideally has 4 or fewer. If an antigen includes only one type of spa domain (*e.g.* only the SpA(A) or SpA(D) domain), it may include more than one copy of this domain *e.g.* multiple SpA(D) domains in a single polypeptide chain.

[0136] An individual domain within the antigen may be mutated at 1, 2, 3, 4, 5, 6, 7, 8, 9, 10 or more amino acids relative to SEQ ID NO: 36 (*e.g.* see ref. 71, disclosing mutations at residues 3 and/or 24 of domain D, at residue 46 and/or 53 of domain A, *etc.*). Such mutants should not remove the antigen's ability to elicit an antibody that recognises SEQ ID NO: 36, but may remove the antigen's binding to IgG and/or other human proteins (such as human blood proteins).

[0137] In certain aspects a spa antigen includes a substitution at (a) one or more amino acid substitution in an IgG Fc binding sub-domain of SpA domain A, B, C, D and/or E that disrupts or decreases binding to IgG Fc, and (b) one or more amino acid substitution in a V_H3 binding sub-domain of SpA domain A, B, C, D, and/or E that disrupts or decreases binding to V_H3. In certain embodiments, a variant SpA comprises at least or at most 1, 2, 3, 4, 5, 6, 7, 8, 9, 10 or more variant SpA domain D peptides.

Second antigen group

sta001

[0138] The 'sta001' antigen is annotated as '5'-nucleotidase family protein'. In the NCTC 8325 strain sta001 is SAOUHSC_00025 and has amino acid sequence SEQ ID NO: 37 (GI:88193846). In the Newman strain it is nwmn_0022 (GI:151220234). It has also been referred to as AdsA and SasH and SA0024.

[0139] Useful sta001 antigens can elicit an antibody (*e.g.* when administered to a human) that recognises SEQ ID NO: 37 and/or may comprise an amino acid sequence: (a) having 50% or more identity (*e.g.* 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 37; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 37, wherein 'n' is 7 or more (*e.g.* 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta001 proteins include variants of SEQ ID NO: 37. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 37. Other preferred fragments lack one or more amino acids (*e.g.* 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (*e.g.* 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 37 while retaining at least one epitope of SEQ ID NO: 37. The final 34 C-terminal amino acids of SEQ ID NO: 37 can usefully be omitted. The first 38 N-terminal amino acids of SEQ ID NO: 37 can usefully be omitted. Other fragments omit one or more protein domains.

sta002

[0140] The 'sta002' antigen is annotated as 'lipoprotein'. In the NCTC 8325 strain sta002 is SAOUHSC_00356 and has amino acid sequence SEQ ID NO: 38 (GI:88194155). In the Newman strain it is nwmn_0364 (GI:151220576).

[0141] Useful sta002 antigens can elicit an antibody (*e.g.* when administered to a human) that recognises SEQ ID NO: 38 and/or may comprise an amino acid sequence: (a) having 50% or more identity (*e.g.* 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 38; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 38, wherein 'n' is 7 or more (*e.g.* 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150 or more). These sta002 proteins include variants of SEQ ID NO: 38. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 38. Other preferred fragments lack one or more amino acids (*e.g.* 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (*e.g.* 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 38 while retaining at least one epitope of SEQ ID NO: 38. The first 18 N-terminal amino acids of SEQ ID NO: 38 can usefully be omitted. Other fragments omit one or more protein domains.

[0142] SEQ ID NOs: 153 ('sta002₁₉₋₁₈₇') and 154 ('sta002₁₉₋₁₂₄') are two useful fragments of SEQ ID NO: 38 which reduce the antigen's similarity with human proteins.

sta003

[0143] The 'sta003' antigen is annotated as 'surface protein'. In the NCTC 8325 strain sta003 is SAOUHSC_00400 and has amino acid sequence SEQ ID NO: 39 (GI:88194195). In the Newman strain it is nwmn_0401 (GI:151220613).

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[0144] Useful sta003 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 39 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 39; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 39, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta003 proteins include variants of SEQ ID NO: 39. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 39. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 39 while retaining at least one epitope of SEQ ID NO: 39. The first 32 N-terminal amino acids of SEQ ID NO: 39 can usefully be omitted. Other fragments omit one or more protein domains.

sta004

[0145] The 'sta004' antigen is annotated as 'Siderophore binding protein FatB'. In the NCTC 8325 strain sta004 is SAOUHSC_00749 and has amino acid sequence SEQ ID NO: 40 (GI:88194514). In the Newman strain it is nwmn_0705 (GI: 151220917).

[0146] Useful sta004 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 40 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 40; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 40, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta004 proteins include variants of SEQ ID NO: 40. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 40. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 40 while retaining at least one epitope of SEQ ID NO: 40. The first 18 N-terminal amino acids of SEQ ID NO: 40 can usefully be omitted. Other fragments omit one or more protein domains.

sta005

[0147] The 'sta005' antigen is annotated as 'superantigen-like protein'. In the NCTC 8325 strain sta005 is SAOUHSC_01127 and has amino acid sequence SEQ ID NO: 41 (GI:88194870). In the Newman strain it is nwmn_1077 (GI: 151221289).

[0148] Useful sta005 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 41 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 41; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 41, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta005 proteins include variants of SEQ ID NO: 41. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 41. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 41 while retaining at least one epitope of SEQ ID NO: 41. The first 18 N-terminal amino acids of SEQ ID NO: 41 can usefully be omitted. Other fragments omit one or more protein domains.

sta006

[0149] The 'sta006' antigen is annotated as 'ferrichrome-binding protein', and has also been referred to as 'FhuD2' in the literature [72]. In the NCTC 8325 strain sta006 is SAOUHSC_02554 and has amino acid sequence SEQ ID NO: 42 (GI:88196199). In the Newman strain it is nwmn_2185 (GI: 151222397).

[0150] Useful sta006 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 42 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 42; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 42, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta006 proteins include variants of SEQ ID NO: 42. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 42. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 42 while retaining at least one epitope of SEQ ID NO: 42. The first 17 N-terminal amino acids of SEQ ID NO: 42 can usefully be omitted (to provide SEQ ID NO: 246). Other fragments omit one or more protein domains. Mutant forms of sta006 are reported

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in reference 73. A sta006 antigen may be lipidated e.g. with an acylated N-terminus cysteine. One useful sta006 sequence is SEQ ID NO: 248, which has a Met-Ala-Ser- sequence at the N-terminus.

sta007

[0151] The 'sta007' antigen is annotated as 'secretory antigen precursor'. In the NCTC 8325 strain sta007 is SAOUHSC_02571 and has amino acid sequence SEQ ID NO: 43 (GI:88196215). In the Newman strain it is nwmn_2199 (GI:151222411). Proteomic analysis has revealed that this protein is secreted or surface-exposed.

[0152] Useful sta007 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 43 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 43; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 43, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta007 proteins include variants of SEQ ID NO: 43. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 43. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 43 while retaining at least one epitope of SEQ ID NO: 43. The first 27 N-terminal amino acids of SEQ ID NO: 43 can usefully be omitted. Other fragments omit one or more protein domains.

sta008

[0153] The 'sta008' antigen is annotated as 'lipoprotein'. In the NCTC 8325 strain sta008 is SAOUHSC_02650 and has amino acid sequence SEQ ID NO: 44 (GI:88196290). In the Newman strain it is nwmn_2270 (GI:151222482).

[0154] Useful sta008 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 44 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 44; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 44, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta008 proteins include variants of SEQ ID NO: 44. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 44. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 44 while retaining at least one epitope of SEQ ID NO: 44. The first 17 N-terminal amino acids of SEQ ID NO: 44 can usefully be omitted. Other fragments omit one or more protein domains.

sta009

[0155] The 'sta009' antigen is annotated as 'immunoglobulin G-binding protein Sbi'. In the NCTC 8325 strain sta009 is SAOUHSC_02706 and has amino acid sequence SEQ ID NO: 45 (GI:88196346). In the Newman strain it is nwmn_2317 (GI: 151222529).

[0156] Useful sta009 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 45 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 45; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 45, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta009 proteins include variants of SEQ ID NO: 45. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 45. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 45 while retaining at least one epitope of SEQ ID NO: 45. The first 29 N-terminal amino acids of SEQ ID NO: 45 can usefully be omitted. Other fragments omit one or more protein domains.

sta010

[0157] The 'sta010' antigen is annotated as 'immunodominant antigen A'. In the NCTC 8325 strain sta010 is SAOUHSC_02887 and has amino acid sequence SEQ ID NO: 46 (GI:88196515). In the Newman strain it is nwmn_2469 (GI:151222681). Proteomic analysis has revealed that this protein is secreted or surface-exposed.

[0158] Useful sta010 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 46 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 46; and/or (b)

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comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 46, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta010 proteins include variants of SEQ ID NO: 46. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 46. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 46 while retaining at least one epitope of SEQ ID NO: 46. The first 29 N-terminal amino acids of SEQ ID NO: 46 can usefully be omitted. Other fragments omit one or more protein domains.

sta011

[0159] The 'sta011' antigen is annotated as 'lipoprotein'. In the NCTC 8325 strain sta011 is SAOUHSC_00052 and has amino acid sequence SEQ ID NO: 47 (GI:88193872).

[0160] Useful sta011 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 47 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 47; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 47, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta011 proteins include variants of SEQ ID NO: 47. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 47. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 47 while retaining at least one epitope of SEQ ID NO: 47. The first 23 N-terminal amino acids of SEQ ID NO: 47 can usefully be omitted (to provide SEQ ID NO: 247). Other fragments omit one or more protein domains. A sta011 antigen may be lipidated e.g. with an acylated N-terminus cysteine. One useful sta011 sequence is SEQ ID NO: 249, which has a N-terminus methionine.

[0161] Variant forms of SEQ ID NO: 47 which may be used as or for preparing sta011 antigens include, but are not limited to, SEQ ID NOs: 213, 214 and 215 with various Ile/Val/Leu substitutions.

[0162] Sta011 can exist as a monomer or an oligomer, with Ca⁺⁺ ions favouring oligomerisation. The invention can use monomers and/or oligomers of Sta011.

sta012

[0163] The 'sta012' antigen is annotated as 'protein with leader'. In the NCTC 8325 strain sta012 is SAOUHSC_00106 and has amino acid sequence SEQ ID NO: 48 (GI:88193919).

[0164] Useful sta012 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 48 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 48; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 48, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta012 proteins include variants of SEQ ID NO: 48. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 48. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 48 while retaining at least one epitope of SEQ ID NO: 48. The first 21 N-terminal amino acids of SEQ ID NO: 48 can usefully be omitted. Other fragments omit one or more protein domains.

sta013

[0165] The 'sta013' antigen is annotated as 'poly-gamma-glutamate capsule biosynthesis protein'. In the NCTC 8325 strain sta013 is SAOUHSC_00107 and has amino acid sequence SEQ ID NO: 49 (GI:88193920).

[0166] Useful sta013 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 49 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 49; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 49, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta013 proteins include variants of SEQ ID NO: 49. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 49. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 49 while retaining at least one epitope of SEQ ID NO: 49. Other fragments omit one or more protein domains.

sta014

[0167] The 'sta014' antigen is annotated as 'lipoprotein'. In the NCTC 8325 strain sta014 is SAOUHSC_00137 and has amino acid sequence SEQ ID NO: 50 (GI:88193950).

[0168] Useful sta014 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 50 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 50; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 50, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta014 proteins include variants of SEQ ID NO: 50. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 50. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 50 while retaining at least one epitope of SEQ ID NO: 50. The first 17 N-terminal amino acids of SEQ ID NO: 50 can usefully be omitted. Other fragments omit one or more protein domains.

sta015

[0169] The 'sta015' antigen is annotated as 'extracellular solute-binding protein; RGD containing lipoprotein'. In the NCTC 8325 strain sta015 is SAOUHSC_00170 and has amino acid sequence SEQ ID NO: 51 (GI:88193980).

[0170] Useful sta015 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 51 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 51; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 51, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta015 proteins include variants of SEQ ID NO: 51. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 51. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 51 while retaining at least one epitope of SEQ ID NO: 51. The first 18 N-terminal amino acids of SEQ ID NO: 51 can usefully be omitted. Other fragments omit one or more protein domains.

sta016

[0171] The 'sta016' antigen is annotated as 'gamma-glutamyltranspeptidase'. In the NCTC 8325 strain sta016 is SAOUHSC_00171 and has amino acid sequence SEQ ID NO: 52 (GI:88193981).

[0172] Useful sta016 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 52 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 52; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 52, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta016 proteins include variants of SEQ ID NO: 52. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 52. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 52 while retaining at least one epitope of SEQ ID NO: 52. Other fragments omit one or more protein domains.

sta017

[0173] The 'sta017' antigen is annotated as 'lipoprotein'. In the NCTC 8325 strain sta017 is SAOUHSC_00186 and has amino acid sequence SEQ ID NO: 53 (GI:88193996).

[0174] Useful sta017 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 53 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 53; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 53, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta017 proteins include variants of SEQ ID NO: 53. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 53. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 53 while retaining at least one epitope of SEQ ID NO: 53. The first 17 N-terminal amino acids of SEQ ID NO: 53 can usefully be omitted. Other fragments omit one or more protein domains.

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sta018

[0175] The 'sta018' antigen is annotated as 'extracellular solute-binding protein'. In the NCTC 8325 strain sta018 is SAOUHSC_00201 and has amino acid sequence SEQ ID NO: 54 (GI:88194011).

[0176] Useful sta018 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 54 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 54; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 54, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta018 proteins include variants of SEQ ID NO: 54. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 54. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 54 while retaining at least one epitope of SEQ ID NO: 54. Other fragments omit one or more protein domains.

sta019

[0177] The 'sta019' antigen is annotated as 'peptidoglycan hydrolase'. In the NCTC 8325 strain sta019 is SAOUHSC_00248 and has amino acid sequence SEQ ID NO: 55 (GI:88194055). In the Newman strain it is nwmn_0210 (GI:151220422).

[0178] Useful sta019 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 55 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 55; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 55, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta019 proteins include variants of SEQ ID NO: 55. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 55. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 55 while retaining at least one epitope of SEQ ID NO: 55. The first 25 N-terminal amino acids of SEQ ID NO: 55 can usefully be omitted. Other fragments omit one or more protein domains. Useful fragments are SEQ ID NOs: 228 and 229.

[0179] Sta019 does not adsorb well to aluminium hydroxide adjuvants, so Sta019 present in a composition may be unadsorbed or may be adsorbed to an alternative adjuvant e.g. to an aluminium phosphate.

sta020

[0180] The 'sta020' antigen is annotated as 'exported protein'. In the NCTC 8325 strain sta020 is SAOUHSC_00253 and has amino acid sequence SEQ ID NO: 56 (GI:88194059).

[0181] Useful sta020 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 56 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 56; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 56, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta020 proteins include variants of SEQ ID NO: 56. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 56. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 56 while retaining at least one epitope of SEQ ID NO: 56. The first 30 N-terminal amino acids of SEQ ID NO: 56 can usefully be omitted. Other fragments omit one or more protein domains.

sta021

[0182] The 'sta021' antigen is annotated as 'secretory antigen SsaA-like protein'. In the NCTC 8325 strain sta021 is SAOUHSC_00256 and has amino acid sequence SEQ ID NO: 57 (GI:88194062).

[0183] Useful sta021 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 57 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 57; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 57, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta021 proteins include variants of SEQ ID NO: 57. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 57. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more

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amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 57 while retaining at least one epitope of SEQ ID NO: 57. The first 24 N-terminal amino acids of SEQ ID NO: 57 can usefully be omitted. Other fragments omit one or more protein domains.

5 *sta022*

[0184] The 'sta022' antigen is annotated as 'lipoprotein'. In the NCTC 8325 strain sta022 is SAOUHSC_00279 and has amino acid sequence SEQ ID NO: 58 (GI:88194083).

10 **[0185]** Useful sta022 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 58 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 58; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 58, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100 or more). These sta022 proteins include variants of SEQ ID NO: 58. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 58. Other preferred fragments lack one or
15 more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 58 while retaining at least one epitope of SEQ ID NO: 58. The first 17 N-terminal amino acids of SEQ ID NO: 58 can usefully be omitted. Other fragments omit one or more protein domains.

20 *sta023*

[0186] The 'sta023' antigen is annotated as '5'-nucleotidase; lipoprotein e(P4) family'. In the NCTC 8325 strain sta023 is SAOUHSC_00284 and has amino acid sequence SEQ ID NO: 59 (GI:88194087).

25 **[0187]** Useful sta023 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 59 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 59; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 59, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta023 proteins include variants of SEQ ID NO: 59. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 59. Other preferred fragments lack
30 one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 59 while retaining at least one epitope of SEQ ID NO: 59. The first 31 N-terminal amino acids of SEQ ID NO: 59 can usefully be omitted. Other fragments omit one or more protein domains.

35 *sta024*

[0188] The 'sta024' antigen is annotated as 'lipase precursor'. In the NCTC 8325 strain sta024 is SAOUHSC_00300 and has amino acid sequence SEQ ID NO: 60 (GI:88194101).

40 **[0189]** Useful sta024 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 60 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 60; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 60, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta024 proteins include variants of SEQ ID NO: 60. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 60. Other preferred fragments lack
45 one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 60 while retaining at least one epitope of SEQ ID NO: 60. The first 37 N-terminal amino acids of SEQ ID NO: 60 can usefully be omitted. Other fragments omit one or more protein domains.

50 *sta025*

[0190] The 'sta025' antigen is annotated as 'lipoprotein'. In the NCTC 8325 strain sta025 is SAOUHSC_00362 and has amino acid sequence SEQ ID NO: 61 (GI:88194160).

55 **[0191]** Useful sta025 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 61 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 61; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 61, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta025 proteins include variants of SEQ

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ID NO: 61. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 61. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 61 while retaining at least one epitope of SEQ ID NO: 61. The first 19 N-terminal amino acids of SEQ ID NO: 61 can usefully be omitted. Other fragments omit one or more protein domains.

sta026

[0192] The 'sta026' antigen is annotated as 'lipoprotein'. In the NCTC 8325 strain sta026 is SAOUHSC_00404 and has amino acid sequence SEQ ID NO: 62 (GI:88194198).

[0193] Useful sta026 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 62 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 62; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 62, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta026 proteins include variants of SEQ ID NO: 62. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 62. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 62 while retaining at least one epitope of SEQ ID NO: 62. The first 22 N-terminal amino acids of SEQ ID NO: 62 can usefully be omitted. Other fragments omit one or more protein domains.

sta027

[0194] The 'sta027' antigen is annotated as 'probable lipase'. In the NCTC 8325 strain sta027 is SAOUHSC_00661 and has amino acid sequence SEQ ID NO: 63 (GI:88194426).

[0195] Useful sta027 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 63 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 63; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 63, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta027 proteins include variants of SEQ ID NO: 63. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 63. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 63 while retaining at least one epitope of SEQ ID NO: 63. The first 23 N-terminal amino acids of SEQ ID NO: 63 can usefully be omitted. Other fragments omit one or more protein domains.

sta028

[0196] The 'sta028' antigen is annotated as 'secretory antigen SsaA-like protein'. In the NCTC 8325 strain sta028 is SAOUHSC_00671 and has amino acid sequence SEQ ID NO: 64 (GI:88194436). In the Newman strain it is nwmn_0634 (GI:151220846).

[0197] Useful sta028 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 64 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 64; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 64, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta028 proteins include variants of SEQ ID NO: 64. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 64. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 64 while retaining at least one epitope of SEQ ID NO: 64. The first 25 N-terminal amino acids of SEQ ID NO: 64 can usefully be omitted. Other fragments omit one or more protein domains.

sta029

[0198] The 'sta029' antigen is annotated as 'ferrichrome binding protein'. In the NCTC 8325 strain sta029 is SAOUHSC_00754 and has amino acid sequence SEQ ID NO: 65 (GI:88194518).

[0199] Useful sta029 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 65 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%,

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80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 65; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 65, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta029 proteins include variants of SEQ ID NO: 65. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 65. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 65 while retaining at least one epitope of SEQ ID NO: 65. The final 25 C-terminal amino acids of SEQ ID NO: 65 can usefully be omitted. The first 19 N-terminal amino acids of SEQ ID NO: 65 can usefully be omitted. Other fragments omit one or more protein domains.

sta030

[0200] The 'sta030' antigen is annotated as 'lipoprotein'. In the NCTC 8325 strain sta030 is SAOUHSC_00808 and has amino acid sequence SEQ ID NO: 66 (GI:88194568).

[0201] Useful sta030 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 66 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 66; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 66, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta030 proteins include variants of SEQ ID NO: 66. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 66. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 66 while retaining at least one epitope of SEQ ID NO: 66. The first 17 N-terminal amino acids of SEQ ID NO: 66 can usefully be omitted. Other fragments omit one or more protein domains.

sta031

[0202] The 'sta031' antigen is annotated as '5-nucleotidase family protein'. In the NCTC 8325 strain sta031 is SAOUHSC_00860 and has amino acid sequence SEQ ID NO: 67 (GI:88194617).

[0203] Useful sta031 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 67 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 67; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 67, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta031 proteins include variants of SEQ ID NO: 67. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 67. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 67 while retaining at least one epitope of SEQ ID NO: 67. Other fragments omit one or more protein domains.

sta032

[0204] The 'sta032' antigen is annotated as 'serine protease HtrA'. In the NCTC 8325 strain sta032 is SAOUHSC_00958 and has amino acid sequence SEQ ID NO: 68 (GI:88194715).

[0205] Useful sta032 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 68 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 68; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 68, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta032 proteins include variants of SEQ ID NO: 68. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 68. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 68 while retaining at least one epitope of SEQ ID NO: 68. Other fragments omit one or more protein domains.

sta033

[0206] The 'sta033' antigen is annotated as 'cysteine protease precursor'. In the NCTC 8325 strain sta033 is SAOUHSC_00987 and has amino acid sequence SEQ ID NO: 69 (GI:88194744).

[0207] Useful sta033 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID

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NO: 69 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 69; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 69, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta033 proteins include variants of SEQ ID NO: 69. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 69. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 69 while retaining at least one epitope of SEQ ID NO: 69. The first 29 N-terminal amino acids of SEQ ID NO: 69 can usefully be omitted. Other fragments omit one or more protein domains.

sta034

[0208] The 'sta034' antigen is annotated as 'glutamyl endopeptidase precursor'. In the NCTC 8325 strain sta034 is SAOUHSC_00988 and has amino acid sequence SEQ ID NO: 70 (GI:88194745).

[0209] Useful sta034 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 70 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 70; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 70, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta034 proteins include variants of SEQ ID NO: 70. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 70. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 70 while retaining at least one epitope of SEQ ID NO: 70. The first 29 N-terminal amino acids of SEQ ID NO: 70 can usefully be omitted. Other fragments omit one or more protein domains.

sta035

[0210] The 'sta035' antigen is annotated as 'fmt protein'. In the NCTC 8325 strain sta035 is SAOUHSC_00998 and has amino acid sequence SEQ ID NO: 71 (GI:88194754).

[0211] Useful sta035 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 71 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 71; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 71, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta035 proteins include variants of SEQ ID NO: 71. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 71. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 71 while retaining at least one epitope of SEQ ID NO: 71. The first 25 N-terminal amino acids of SEQ ID NO: 71 can usefully be omitted. Other fragments omit one or more protein domains.

sta036

[0212] The 'sta036' antigen is annotated as 'iron-regulated protein with leader'. In the NCTC 8325 strain sta036 is SAOUHSC_01084 and has amino acid sequence SEQ ID NO: 72 (GI:88194831).

[0213] Useful sta036 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 72 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 72; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 72, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta036 proteins include variants of SEQ ID NO: 72. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 72. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 72 while retaining at least one epitope of SEQ ID NO: 72. The final 27 C-terminal amino acids of SEQ ID NO: 72 can usefully be omitted. The first 32 N-terminal amino acids of SEQ ID NO: 72 can usefully be omitted. Other fragments omit one or more protein domains.

sta037

[0214] The 'sta037' antigen is annotated as 'iron ABC transporter; iron -binding protein lsdE'. In the NCTC 8325 strain sta037 is SAOUHSC_01085 and has amino acid sequence SEQ ID NO: 73 (GI:88194832).

[0215] Useful sta037 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 73 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 73; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 73, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta037 proteins include variants of SEQ ID NO: 73. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 73. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 73 while retaining at least one epitope of SEQ ID NO: 73. The first 9 N-terminal amino acids of SEQ ID NO: 73 can usefully be omitted. Other fragments omit one or more protein domains.

sta038

[0216] The 'sta038' antigen is annotated as 'NPQTN specific sortase B'. In the NCTC 8325 strain sta038 is SAOUHSC_01088 and has amino acid sequence SEQ ID NO: 74 (GI:88194835).

[0217] Useful sta038 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 74 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 74; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 74, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta038 proteins include variants of SEQ ID NO: 74. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 74. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 74 while retaining at least one epitope of SEQ ID NO: 74. The first 21 N-terminal amino acids of SEQ ID NO: 74 can usefully be omitted. Other fragments omit one or more protein domains.

sta039

[0218] The 'sta039' antigen is annotated as 'superantigen-like protein'. In the NCTC 8325 strain sta039 is SAOUHSC_01124 and has amino acid sequence SEQ ID NO: 75 (GI:88194868).

[0219] Useful sta039 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 75 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 75; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 75, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta039 proteins include variants of SEQ ID NO: 75. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 75. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 75 while retaining at least one epitope of SEQ ID NO: 75. The first 22 N-terminal amino acids of SEQ ID NO: 75 can usefully be omitted. Other fragments omit one or more protein domains.

sta040

[0220] The 'sta040' antigen is annotated as 'superantigen-like protein'. In the NCTC 8325 strain sta040 is SAOUHSC_01125 and has amino acid sequence SEQ ID NO: 76 (GI:88194869). In the Newman strain it is nwmn_1076 (GI:151221288).

[0221] Useful sta040 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 76 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 76; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 76, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta040 proteins include variants of SEQ ID NO: 76. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 76. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 76 while retaining

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at least one epitope of SEQ ID NO: 76. The first 21 N-terminal amino acids of SEQ ID NO: 76 can usefully be omitted. Other fragments omit one or more protein domains.

sta041

5 **[0222]** The 'sta041' antigen is annotated as 'fibronectin-binding protein A-related'. In the NCTC 8325 strain sta041 is SAOUHSC_01175 and has amino acid sequence SEQ ID NO: 77 (GI:88194914).

10 **[0223]** Useful sta041 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 77 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 77; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 77, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta041 proteins include variants of SEQ ID NO: 77. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 77. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 77 while retaining at least one epitope of SEQ ID NO: 77. Other fragments omit one or more protein domains.

sta042

20 **[0224]** The 'sta042' antigen is annotated as 'lipoprotein'. In the NCTC 8325 strain sta042 is SAOUHSC_01180 and has amino acid sequence SEQ ID NO: 78 (GI:88194919).

25 **[0225]** Useful sta042 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 78 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 78; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 78, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta042 proteins include variants of SEQ ID NO: 78. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 78. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 78 while retaining at least one epitope of SEQ ID NO: 78. The first 18 N-terminal amino acids of SEQ ID NO: 78 can usefully be omitted. Other fragments omit one or more protein domains.

sta043

35 **[0226]** The 'sta043' antigen is annotated as 'cell wall hydrolase'. In the NCTC 8325 strain sta043 is SAOUHSC_01219 and has amino acid sequence SEQ ID NO: 79 (GI:88194955).

40 **[0227]** Useful sta043 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 79 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 79; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 79, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta043 proteins include variants of SEQ ID NO: 79. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 79. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 79 while retaining at least one epitope of SEQ ID NO: 79. The first 38 N-terminal amino acids of SEQ ID NO: 79 can usefully be omitted. Other fragments omit one or more protein domains.

sta044

50 **[0228]** The 'sta044' antigen is annotated as 'lipoprotein'. In the NCTC 8325 strain sta044 is SAOUHSC_01508 and has amino acid sequence SEQ ID NO: 80 (GI:88195223).

55 **[0229]** Useful sta044 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 80 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 80; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 80, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta044 proteins include variants of SEQ ID NO: 80. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 80. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more

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amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 80 while retaining at least one epitope of SEQ ID NO: 80. The first 17 N-terminal amino acids of SEQ ID NO: 80 can usefully be omitted. Other fragments omit one or more protein domains.

5 *sta045*

[0230] The 'sta045' antigen is annotated as 'lipoprotein'. In the NCTC 8325 strain sta045 is SAOUHSC_01627 and has amino acid sequence SEQ ID NO: 81 (GI:88195337).

10 **[0231]** Useful sta045 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 81 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 81; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 81, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150 or more). These sta045 proteins include variants of SEQ ID NO: 81. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 81. Other preferred fragments lack one or
15 more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 81 while retaining at least one epitope of SEQ ID NO: 81. The first 16 N-terminal amino acids of SEQ ID NO: 81 can usefully be omitted. Other fragments omit one or more protein domains.

20 *sta046*

[0232] The 'sta046' antigen is annotated as 'Excalibur protein'. In the NCTC 8325 strain sta046 is SAOUHSC_01918 and has amino acid sequence SEQ ID NO: 82 (GI:88195613).

25 **[0233]** Useful sta046 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 82 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 82; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 82, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta046 proteins include variants of SEQ ID NO: 82. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 82. Other preferred fragments lack one or
30 more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 82 while retaining at least one epitope of SEQ ID NO: 82. The first 53 N-terminal amino acids of SEQ ID NO: 82 can usefully be omitted. Other fragments omit one or more protein domains.

35 *sta047*

[0234] The 'sta047' antigen is annotated as 'lipoprotein'. In the NCTC 8325 strain sta047 is SAOUHSC_01920 and has amino acid sequence SEQ ID NO: 83 (GI:88195615).

40 **[0235]** Useful sta047 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 83 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 83; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 83, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta047 proteins include variants of SEQ ID NO: 83. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 83. Other preferred fragments lack one or
45 more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 83 while retaining at least one epitope of SEQ ID NO: 83. The first 18 N-terminal amino acids of SEQ ID NO: 83 can usefully be omitted. Other fragments omit one or more protein domains.

50 *sta048*

[0236] The 'sta048' antigen is annotated as 'intracellular serine protease'. In the NCTC 8325 strain sta048 is SAOUHSC_01949 and has amino acid sequence SEQ ID NO: 84 (GI:88195642).

55 **[0237]** Useful sta048 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 84 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 84; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 84, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta048 proteins include variants

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of SEQ ID NO: 84. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 84. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 84 while retaining at least one epitope of SEQ ID NO: 84. The first 27 N-terminal amino acids of SEQ ID NO: 84 can usefully be omitted.

sta049

[0238] The 'sta049' antigen is annotated as 'protein export protein PrsA'. In the NCTC 8325 strain sta049 is SAOUHSC_01972 and has amino acid sequence SEQ ID NO: 85 (GI:88195663). In the Newman strain it is nwmn_1733 (GI:151221945).

[0239] Useful sta049 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 85 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 85; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 85, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta049 proteins include variants of SEQ ID NO: 85. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 85. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 85 while retaining at least one epitope of SEQ ID NO: 85. The first 25 N-terminal amino acids of SEQ ID NO: 85 can usefully be omitted. Other fragments omit one or more protein domains.

sta050

[0240] The 'sta050' antigen is annotated as 'staphopain thiol proteinase'. In the NCTC 8325 strain sta050 is SAOUHSC_02127 and has amino acid sequence SEQ ID NO: 86 (GI:88195808).

[0241] Useful sta050 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 86 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 86; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 86, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta050 proteins include variants of SEQ ID NO: 86. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 86. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 86 while retaining at least one epitope of SEQ ID NO: 86. The first 25 N-terminal amino acids of SEQ ID NO: 86 can usefully be omitted. Other fragments omit one or more protein domains.

sta051

[0242] The 'sta051' antigen is annotated as 'protein with leader'. In the NCTC 8325 strain sta051 is SAOUHSC_02147 and has amino acid sequence SEQ ID NO: 87 (GI:88195827).

[0243] Useful sta051 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 87 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 87; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 87, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta051 proteins include variants of SEQ ID NO: 87. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 87. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 87 while retaining at least one epitope of SEQ ID NO: 87. The first 24 N-terminal amino acids of SEQ ID NO: 87 can usefully be omitted. Other fragments omit one or more protein domains.

sta052

[0244] The 'sta052' antigen is annotated as 'ferric hydroxamate receptor 1'. In the NCTC 8325 strain sta052 is SAOUHSC_02246 and has amino acid sequence SEQ ID NO: 88 (GI:88195918).

[0245] Useful sta052 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 88 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%,

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80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 88; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 88, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta052 proteins include variants of SEQ ID NO: 88. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 88. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 88 while retaining at least one epitope of SEQ ID NO: 88. The first 17 N-terminal amino acids of SEQ ID NO: 88 can usefully be omitted. Other fragments omit one or more protein domains.

10 *sta053*

[0246] The 'sta053' antigen is annotated as 'srdH family protein'. In the NCTC 8325 strain sta053 is SAOUHSC_02257 and has amino acid sequence SEQ ID NO: 89 (GI:88195928).

[0247] Useful sta053 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 89 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 89; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 89, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta053 proteins include variants of SEQ ID NO: 89. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 89. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 89 while retaining at least one epitope of SEQ ID NO: 89. The first 26 N-terminal amino acids of SEQ ID NO: 89 can usefully be omitted. Other fragments omit one or more protein domains.

25 *sta054*

[0248] The 'sta054' antigen is annotated as 'Probable transglycosylase isaA precursor'. In the NCTC 8325 strain sta054 is SAOUHSC_02333 and has amino acid sequence SEQ ID NO: 90 (GI:88195999). Useful sta054 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 90 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 90; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 90, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta054 proteins include variants of SEQ ID NO: 90. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 90. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 90 while retaining at least one epitope of SEQ ID NO: 90. The first 27 N-terminal amino acids of SEQ ID NO: 90 can usefully be omitted. Other fragments omit one or more protein domains.

40 *sta055*

[0249] The 'sta055' antigen is annotated as 'surface hydrolase'. In the NCTC 8325 strain sta055 is SAOUHSC_02448 and has amino acid sequence SEQ ID NO: 91 (GI:88196100).

[0250] Useful sta055 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 91 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 91; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 91, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta055 proteins include variants of SEQ ID NO: 91. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 91. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 91 while retaining at least one epitope of SEQ ID NO: 91. The first 31 N-terminal amino acids of SEQ ID NO: 91 can usefully be omitted. Other fragments omit one or more protein domains.

55 *sta056*

[0251] The 'sta056' antigen is annotated as 'hyaluronate lyase'. In the NCTC 8325 strain sta056 is SAOUHSC_02463 and has amino acid sequence SEQ ID NO: 92 (GI:88196115).

[0252] Useful sta056 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID

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NO: 92 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 92; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 92, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta056 proteins include variants of SEQ ID NO: 92. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 92. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 92 while retaining at least one epitope of SEQ ID NO: 92. The first 24 N-terminal amino acids of SEQ ID NO: 92 can usefully be omitted. Other fragments omit one or more protein domains.

sta057

[0253] The 'sta057' antigen is annotated as 'secretory antigen precursor SsaA'. In the NCTC 8325 strain sta057 is SAOUHSC_02576 and has amino acid sequence SEQ ID NO: 93 (GI:88196220). In the Newman strain it is nwmn_2203 (GI:151222415).

[0254] Useful sta057 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 93 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 93; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 93, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150 or more). These sta057 proteins include variants of SEQ ID NO: 93. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 93. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 93 while retaining at least one epitope of SEQ ID NO: 93. The first 27 N-terminal amino acids of SEQ ID NO: 93 can usefully be omitted. Other fragments omit one or more protein domains.

sta058

[0255] The 'sta058' antigen is annotated as 'Zn-binding lipoprotein adcA-like'. In the NCTC 8325 strain sta058 is SAOUHSC_02690 and has amino acid sequence SEQ ID NO: 94 (GI:88196330).

[0256] Useful sta058 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 94 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 94; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 94, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta058 proteins include variants of SEQ ID NO: 94. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 94. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 94 while retaining at least one epitope of SEQ ID NO: 94. The first 20 N-terminal amino acids of SEQ ID NO: 94 can usefully be omitted. Other fragments omit one or more protein domains.

sta059

[0257] The 'sta059' antigen is annotated as 'gamma-hemolysin h-gamma-ii subunit'. In the NCTC 8325 strain sta059 is SAOUHSC_02708 and has amino acid sequence SEQ ID NO: 95 (GI:88196348).

[0258] Useful sta059 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 95 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 95; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 95, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta059 proteins include variants of SEQ ID NO: 95. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 95. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 95 while retaining at least one epitope of SEQ ID NO: 95. The first 20 N-terminal amino acids of SEQ ID NO: 95 can usefully be omitted. Other fragments omit one or more protein domains.

sta060

[0259] The 'sta060' antigen is annotated as 'peptide ABC transporter; peptide-binding protein'. In the NCTC 8325 strain sta060 is SAOUHSC_02767 and has amino acid sequence SEQ ID NO: 96 (GI:88196403).

[0260] Useful sta060 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 96 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 96; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 96, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta060 proteins include variants of SEQ ID NO: 96. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 96. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 96 while retaining at least one epitope of SEQ ID NO: 96. The first 20 N-terminal amino acids of SEQ ID NO: 96 can usefully be omitted. Other fragments omit one or more protein domains.

sta061

[0261] The 'sta061' antigen is annotated as 'protein with leader'. In the NCTC 8325 strain sta061 is SAOUHSC_02783 and has amino acid sequence SEQ ID NO: 97 (GI:88196419).

[0262] Useful sta061 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 97 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 97; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 97, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta061 proteins include variants of SEQ ID NO: 97. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 97. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 97 while retaining at least one epitope of SEQ ID NO: 97. The first 21 N-terminal amino acids of SEQ ID NO: 97 can usefully be omitted. Other fragments omit one or more protein domains.

sta062

[0263] The 'sta062' antigen is annotated as 'protein with leader'. In the NCTC 8325 strain sta062 is SAOUHSC_02788 and has amino acid sequence SEQ ID NO: 98 (GI:88196424).

[0264] Useful sta062 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 98 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 98; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 98, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta062 proteins include variants of SEQ ID NO: 98. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 98. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 98 while retaining at least one epitope of SEQ ID NO: 98. The first 22 N-terminal amino acids of SEQ ID NO: 98 can usefully be omitted. Other fragments omit one or more protein domains.

sta063

[0265] The 'sta063' antigen is annotated as 'aureolysin'. In the NCTC 8325 strain sta063 is SAOUHSC_02971 and has amino acid sequence SEQ ID NO: 99 (GI:88196592).

[0266] Useful sta063 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 99 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 99; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 99, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta063 proteins include variants of SEQ ID NO: 99. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 99. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 99 while retaining at least one epitope of SEQ ID NO: 99. The first 16 N-terminal amino acids of SEQ ID NO: 99 can usefully be omitted.

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Other fragments omit one or more protein domains.

sta064

- 5 **[0267]** The 'sta064' antigen is annotated as 'lipase'. In the NCTC 8325 strain sta064 is SAOUHSC_03006 and has amino acid sequence SEQ ID NO: 100 (GI:88196625). In the Newman strain it is nwmn_2569 (GI:151222781).
- [0268]** Useful sta064 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 100 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 100; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 100, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta064 proteins include variants of SEQ ID NO: 100. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 100. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 100 while retaining at least one epitope of SEQ ID NO: 100. The first 34 N-terminal amino acids of SEQ ID NO: 100 can usefully be omitted. Other fragments omit one or more protein domains.

sta065

- 20 **[0269]** The 'sta065' antigen is annotated as '1-phosphatidylinositol phosphodiesterase precursor'. In the NCTC 8325 strain sta065 is SAOUHSC_00051 and has amino acid sequence SEQ ID NO: 101 (GI:88193871).
- [0270]** Useful sta065 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 101 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 101; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 101, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta065 proteins include variants of SEQ ID NO: 101. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 101. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 101 while retaining at least one epitope of SEQ ID NO: 101. The first 26 N-terminal amino acids of SEQ ID NO: 101 can usefully be omitted. Other fragments omit one or more protein domains.

sta066

- 35 **[0271]** The 'sta066' antigen is annotated as 'protein'. In the NCTC 8325 strain sta066 is SAOUHSC_00172 and has amino acid sequence SEQ ID NO: 102 (GI:88193982).
- [0272]** Useful sta066 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 102 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 102; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 102, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta066 proteins include variants of SEQ ID NO: 102. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 102. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 102 while retaining at least one epitope of SEQ ID NO: 102. The first 21 N-terminal amino acids of SEQ ID NO: 102 can usefully be omitted. Other fragments omit one or more protein domains.

sta067

- 50 **[0273]** The 'sta067' antigen is annotated as 'bacterial extracellular solute-binding protein'. In the NCTC 8325 strain sta067 is SAOUHSC_00176 and has amino acid sequence SEQ ID NO: 103 (GI:88193986).
- [0274]** Useful sta067 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 103 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 103; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 103, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta067 proteins include variants of SEQ ID NO: 103. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 103. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or

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more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 103 while retaining at least one epitope of SEQ ID NO: 103. The first 20 N-terminal amino acids of SEQ ID NO: 103 can usefully be omitted. Other fragments omit one or more protein domains.

5 *sta068*

[0275] The 'sta068' antigen is annotated as 'iron permease FTR1'. In the NCTC 8325 strain sta068 is SAOUHSC_00327 and has amino acid sequence SEQ ID NO: 104 (GI:88194127).

10 **[0276]** Useful sta068 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 104 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 104; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 104, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta068 proteins include variants of SEQ ID NO: 104. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 104. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 104 while retaining at least one epitope of SEQ ID NO: 104. The final 20 C-terminal amino acids of SEQ ID NO: 104 can usefully be omitted. The first 14 N-terminal amino acids of SEQ ID NO: 104 can usefully be omitted. Other fragments omit one or more protein domains.

20

sta069

[0277] The 'sta069' antigen is annotated as 'autolysin precursor'. In the NCTC 8325 strain sta069 is SAOUHSC_00427 and has amino acid sequence SEQ ID NO: 105 (GI:88194219).

25 **[0278]** Useful sta069 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 105 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 105; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 105, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta069 proteins include variants of SEQ ID NO: 105. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 105. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 105 while retaining at least one epitope of SEQ ID NO: 105. The first 25 N-terminal amino acids of SEQ ID NO: 105 can usefully be omitted. Other fragments omit one or more protein domains.

35

sta070

[0279] The 'sta070' antigen is annotated as 'immunogenic secreted precursor-like protein (truncated)'. In the NCTC 8325 strain sta070 is SAOUHSC_00773 and has amino acid sequence SEQ ID NO: 106 (GI:88194535).

40 **[0280]** Useful sta070 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 106 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 106; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 106, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta070 proteins include variants of SEQ ID NO: 106. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 106. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 106 while retaining at least one epitope of SEQ ID NO: 106. The first 24 N-terminal amino acids of SEQ ID NO: 106 can usefully be omitted. Other fragments omit one or more protein domains.

50

sta071

[0281] The 'sta071' antigen is annotated as 'hemolysin'. In the NCTC 8325 strain sta071 is SAOUHSC_00854 and has amino acid sequence SEQ ID NO: 107 (GI:88194612).

55 **[0282]** Useful sta071 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 107 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 107; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 107, wherein 'n' is 7 or more (e.g. 8, 10,

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12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta071 proteins include variants of SEQ ID NO: 107. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 107. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 107 while retaining at least one epitope of SEQ ID NO: 107. The first 24 N-terminal amino acids of SEQ ID NO: 107 can usefully be omitted. Other fragments omit one or more protein domains.

sta072

[0283] The 'sta072' antigen is annotated as 'extramembranal protein'. In the NCTC 8325 strain sta072 is SAOUHSC_00872 and has amino acid sequence SEQ ID NO: 108 (GI:88194629).

[0284] Useful sta072 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 108 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 108; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 108, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta072 proteins include variants of SEQ ID NO: 108. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 108. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 108 while retaining at least one epitope of SEQ ID NO: 108. The first 24 N-terminal amino acids of SEQ ID NO: 108 can usefully be omitted. Other fragments omit one or more protein domains.

sta073

[0285] The 'sta073' antigen is annotated as 'bifunctional autolysin precursor'. In the NCTC 8325 strain sta073 is SAOUHSC_00994 and has amino acid sequence SEQ ID NO: 109 (GI:88194750). In the Newman strain it is nwmn_0922 (GI: 151221134). Proteomic analysis has revealed that this protein is secreted or surface-exposed.

[0286] Useful sta073 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 109 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 109; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 109, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta073 proteins include variants of SEQ ID NO: 109. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 109. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 109 while retaining at least one epitope of SEQ ID NO: 109. The first 24 N-terminal amino acids of SEQ ID NO: 109 can usefully be omitted. Other fragments omit one or more protein domains.

[0287] A Sta073 antigen can usefully be included in a composition in combination with a Sta112 [74].

[0288] Sta073 does not adsorb well to aluminium hydroxide adjuvants, so Sta073 present in a composition may be unadsorbed or may be adsorbed to an alternative adjuvant e.g. to an aluminium phosphate.

sta074

[0289] The 'sta074' antigen is annotated as 'factor essential for methicillin resistance'. In the NCTC 8325 strain sta074 is SAOUHSC_01220 and has amino acid sequence SEQ ID NO: 110 (GI:88194956).

[0290] Useful sta074 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 110 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 110; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 110, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta074 proteins include variants of SEQ ID NO: 110. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 110. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 110 while retaining at least one epitope of SEQ ID NO: 110. Other fragments omit one or more protein domains.

sta075

[0291] The 'sta075' antigen is annotated as 'insulysin; peptidase family M16'. In the NCTC 8325 strain sta075 is

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SAOUHSC_01256 and has amino acid sequence SEQ ID NO: 111 (GI:88194989).

[0292] Useful sta075 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 111 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 111; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 111, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta075 proteins include variants of SEQ ID NO: 111. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 111. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 111 while retaining at least one epitope of SEQ ID NO: 111. Other fragments omit one or more protein domains.

sta076

[0293] The 'sta076' antigen is annotated as 'hydrolase'. In the NCTC 8325 strain sta076 is SAOUHSC_01263 and has amino acid sequence SEQ ID NO: 112 (GI:88194996).

[0294] Useful sta076 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 112 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 112; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 112, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta076 proteins include variants of SEQ ID NO: 112. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 112. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 112 while retaining at least one epitope of SEQ ID NO: 112. The first 24 N-terminal amino acids of SEQ ID NO: 112 can usefully be omitted. Other fragments omit one or more protein domains.

sta077

[0295] The 'sta077' antigen is annotated as 'protein'. In the NCTC 8325 strain sta077 is SAOUHSC_01317 and has amino acid sequence SEQ ID NO: 113 (GI:88195047). Proteomic analysis has revealed that this protein is secreted or surface-exposed.

[0296] Useful sta077 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 113 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 113; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 113, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta077 proteins include variants of SEQ ID NO: 113. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 113. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 113 while retaining at least one epitope of SEQ ID NO: 113. The first 20 N-terminal amino acids of SEQ ID NO: 113 can usefully be omitted. Other fragments omit one or more protein domains.

sta078

[0297] The 'sta078' antigen is annotated as 'FtsK/SpoIIIE family protein'. In the NCTC 8325 strain sta078 is SAOUHSC_01857 and has amino acid sequence SEQ ID NO: 114 (GI:88195555).

[0298] Useful sta078 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 114 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 114; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 114, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta078 proteins include variants of SEQ ID NO: 114. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 114. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 114 while retaining at least one epitope of SEQ ID NO: 114. Other fragments omit one or more protein domains.

sta079

[0299] The 'sta079' antigen is annotated as 'serine protease SpIF'. In the NCTC 8325 strain sta079 is SAOUHSC_01935 and has amino acid sequence SEQ ID NO: 115 (GI:88195630).

[0300] Useful sta079 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 115 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 115; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 115, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta079 proteins include variants of SEQ ID NO: 115. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 115. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 115 while retaining at least one epitope of SEQ ID NO: 115. The first 36 N-terminal amino acids of SEQ ID NO: 115 can usefully be omitted. Other fragments omit one or more protein domains.

sta080

[0301] The 'sta080' antigen is annotated as 'serine protease SpIE'. In the NCTC 8325 strain sta080 is SAOUHSC_01936 and has amino acid sequence SEQ ID NO: 116 (GI:88195631).

[0302] Useful sta080 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 116 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 116; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 116, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta080 proteins include variants of SEQ ID NO: 116. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 116. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 116 while retaining at least one epitope of SEQ ID NO: 116. The first 36 N-terminal amino acids of SEQ ID NO: 116 can usefully be omitted. Other fragments omit one or more protein domains.

sta081

[0303] The 'sta081' antigen is annotated as 'serine protease SpID (EC:3.4.21.19)'. In the NCTC 8325 strain sta081 is SAOUHSC_01938 and has amino acid sequence SEQ ID NO: 170 (GI:88195633).

[0304] Useful sta081 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 170 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 170; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 170, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta081 proteins include variants of SEQ ID NO: 170. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 170. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 170 while retaining at least one epitope of SEQ ID NO: 170. The first 36 N-terminal amino acids of SEQ ID NO: 170 can usefully be omitted. Other fragments omit one or more protein domains.

sta082

[0305] The 'sta082' antigen is annotated as 'serine protease SpIC'. In the NCTC 8325 strain sta082 is SAOUHSC_01939 and has amino acid sequence SEQ ID NO: 117 (GI:88195634).

[0306] Useful sta082 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 117 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 117; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 117, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta082 proteins include variants of SEQ ID NO: 117. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 117. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 117 while retaining at least one epitope of SEQ ID NO: 117. The first 36 N-terminal amino acids of SEQ ID NO: 117 can usefully be omitted.

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Other fragments omit one or more protein domains.

sta083

5 **[0307]** The 'sta083' antigen is annotated as 'serine protease SpIB'. In the NCTC 8325 strain sta083 is SAOUHSC_01941 and has amino acid sequence SEQ ID NO: 118 (GI:88195635).

[0308] Useful sta083 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 118 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 118; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 118, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta083 proteins include variants of SEQ ID NO: 118. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 118. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 118 while retaining at least one epitope of SEQ ID NO: 118. The first 36 N-terminal amino acids of SEQ ID NO: 118 can usefully be omitted. Other fragments omit one or more protein domains.

sta084

20 **[0309]** The 'sta084' antigen is annotated as 'serine protease SpIA'. In the NCTC 8325 strain sta084 is SAOUHSC_01942 and has amino acid sequence SEQ ID NO: 119 (GI:88195636).

[0310] Useful sta084 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 119 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 119; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 119, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta084 proteins include variants of SEQ ID NO: 119. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 119. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 119 while retaining at least one epitope of SEQ ID NO: 119. The first 35 N-terminal amino acids of SEQ ID NO: 119 can usefully be omitted. Other fragments omit one or more protein domains.

sta085

35 **[0311]** The 'sta085' antigen is annotated as 'staphylokinase precursor'. In the NCTC 8325 strain sta085 is SAOUHSC_02171 and has amino acid sequence SEQ ID NO: 120 (GI:88195848).

[0312] Useful sta085 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 120 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 120; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 120, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150 or more). These sta085 proteins include variants of SEQ ID NO: 120. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 120. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 120 while retaining at least one epitope of SEQ ID NO: 120. The first 27 N-terminal amino acids of SEQ ID NO: 120 can usefully be omitted. Other fragments omit one or more protein domains.

sta086

50 **[0313]** The 'sta086' antigen is annotated as 'OxaA-like protein'. In the NCTC 8325 strain sta086 is SAOUHSC_02327 and has amino acid sequence SEQ ID NO: 121 (GI:88195993).

[0314] Useful sta086 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 121 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 121; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 121, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta086 proteins include variants of SEQ ID NO: 121. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 121. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or

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more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 121 while retaining at least one epitope of SEQ ID NO: 121. The first 19 N-terminal amino acids of SEQ ID NO: 121 can usefully be omitted. Other fragments omit one or more protein domains.

5 *sta087*

[0315] The 'sta087' antigen is annotated as 'teicoplanin resistance protein TcaA'. In the NCTC 8325 strain sta087 is SAOUHSC_02635 and has amino acid sequence SEQ ID NO: 122 (GI:88196276).

10 **[0316]** Useful sta087 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 122 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 122; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 122, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta087 proteins include variants of SEQ ID NO: 122. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 122. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 122 while retaining at least one epitope of SEQ ID NO: 122. Other fragments omit one or more protein domains.

20 *sta088*

[0317] The 'sta088' antigen is annotated as 'esterase'. In the NCTC 8325 strain sta088 is SAOUHSC_02844 and has amino acid sequence SEQ ID NO: 123 (GI:88196477).

25 **[0318]** Useful sta088 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 123 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 123; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 123, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta088 proteins include variants of SEQ ID NO: 123. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 123. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 123 while retaining at least one epitope of SEQ ID NO: 123. The first 18 N-terminal amino acids of SEQ ID NO: 123 can usefully be omitted. Other fragments omit one or more protein domains.

35 *sta089*

[0319] The 'sta089' antigen is annotated as 'LysM domain protein'. In the NCTC 8325 strain sta089 is SAOUHSC_02855 and has amino acid sequence SEQ ID NO: 124 (GI:88196486).

40 **[0320]** Useful sta089 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 124 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 124; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 124, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100 or more). These sta089 proteins include variants of SEQ ID NO: 124. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 124. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 124 while retaining at least one epitope of SEQ ID NO: 124. The first 20 N-terminal amino acids of SEQ ID NO: 124 can usefully be omitted. Other fragments omit one or more protein domains.

50 *sta090*

[0321] The 'sta090' antigen is annotated as 'LysM domain protein'. In the NCTC 8325 strain sta090 is SAOUHSC_02883 and has amino acid sequence SEQ ID NO: 125 (GI:88196512).

55 **[0322]** Useful sta090 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 125 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 125; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 125, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta090 proteins include variants of SEQ ID NO: 125. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 125. Other preferred fragments

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lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 125 while retaining at least one epitope of SEQ ID NO: 125. The first 26 N-terminal amino acids of SEQ ID NO: 125 can usefully be omitted. Other fragments omit one or more protein domains.

5

sta091

[0323] The 'sta091' antigen is annotated as 'lipoprotein'. In the NCTC 8325 strain sta091 is SAOUHSC_00685 and has amino acid sequence SEQ ID NO: 126 (GI:88194450).

10 **[0324]** Useful sta091 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 126 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 126; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 126, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100 or more). These sta091 proteins include variants of SEQ ID
15 NO: 126. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 126. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 126 while retaining at least one epitope of SEQ ID NO: 126. The first 15 N-terminal amino acids of SEQ ID NO: 126 can usefully be omitted. Other fragments omit one or more protein domains.

20

sta092

[0325] The 'sta092' antigen is annotated as 'M23/M37 peptidase domain protein'. In the NCTC 8325 strain sta092 is SAOUHSC_00174 and has amino acid sequence SEQ ID NO: 127 (GI:88193984).

25 **[0326]** Useful sta092 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 127 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 127; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 127, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150 or more). These sta092 proteins include variants of SEQ
30 ID NO: 127. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 127. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 127 while retaining at least one epitope of SEQ ID NO: 127. The first 25 N-terminal amino acids of SEQ ID NO: 127 can usefully be omitted. Other fragments omit one or more protein domains.

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sta093

[0327] The 'sta093' antigen is annotated as 'protein'. In the NCTC 8325 strain sta093 is SAOUHSC_01854 and has amino acid sequence SEQ ID NO: 128 (GI:88195552).

40 **[0328]** Useful sta093 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 128 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 128; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 128, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta093 proteins include variants
45 of SEQ ID NO: 128. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 128. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 128 while retaining at least one epitope of SEQ ID NO: 128. Other fragments omit one or more protein domains.

50

sta094

[0329] The 'sta094' antigen is annotated as 'protein'. In the NCTC 8325 strain sta094 is SAOUHSC_01512 and has amino acid sequence SEQ ID NO: 129 (GI:88195226).

55 **[0330]** Useful sta094 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 129 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 129; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 129, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta094 proteins include variants

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of SEQ ID NO: 129. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 129. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 129 while retaining at least one epitope of SEQ ID NO: 129. The first 17 N-terminal amino acids of SEQ ID NO: 129 can usefully be omitted. Other fragments omit one or more protein domains.

sta095

[0331] The 'sta095' antigen is annotated as 'superantigen-like protein'. In the NCTC 8325 strain sta095 is SAOUHSC_00383 and has amino acid sequence SEQ ID NO: 130 (GI:88194180). In the Newman strain it is nwmn_0388 (GI:151220600).

[0332] Useful sta095 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 130 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 130; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 130, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta095 proteins include variants of SEQ ID NO: 130. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 130. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 130 while retaining at least one epitope of SEQ ID NO: 130. The first 32 N-terminal amino acids of SEQ ID NO: 130 can usefully be omitted. Other fragments omit one or more protein domains.

sta096

[0333] The 'sta096' antigen is annotated as 'superantigen-like protein'. In the NCTC 8325 strain sta096 is SAOUHSC_00384 and has amino acid sequence SEQ ID NO: 131 (GI:88194181).

[0334] Useful sta096 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 131 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 131; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 131, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta096 proteins include variants of SEQ ID NO: 131. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 131. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 131 while retaining at least one epitope of SEQ ID NO: 131. The first 30 N-terminal amino acids of SEQ ID NO: 131 can usefully be omitted. Other fragments omit one or more protein domains.

sta097

[0335] The 'sta097' antigen is annotated as 'superantigen-like protein'. In the NCTC 8325 strain sta097 is SAOUHSC_00386 and has amino acid sequence SEQ ID NO: 132 (GI:88194182).

[0336] Useful sta097 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 132 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 132; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 132, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta097 proteins include variants of SEQ ID NO: 132. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 132. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 132 while retaining at least one epitope of SEQ ID NO: 132. The first 30 N-terminal amino acids of SEQ ID NO: 132 can usefully be omitted. Other fragments omit one or more protein domains.

sta098

[0337] The 'sta098' antigen is annotated as 'superantigen-like protein'. In the NCTC 8325 strain sta098 is SAOUHSC_00389 and has amino acid sequence SEQ ID NO: 133 (GI:88194184). In the Newman strain it is nwmn_0391 (GI:151220603).

[0338] Useful sta098 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID

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NO: 133 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 133; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 133, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta098 proteins include variants of SEQ ID NO: 133. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 133. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 133 while retaining at least one epitope of SEQ ID NO: 133. The first 30 N-terminal amino acids of SEQ ID NO: 133 can usefully be omitted. Other fragments omit one or more protein domains.

sta099

[0339] The 'sta099' antigen is annotated as 'superantigen-like protein 5'. In the NCTC 8325 strain sta099 is SAOUHSC_00390 and has amino acid sequence SEQ ID NO: 134 (GI:88194185).

[0340] Useful sta099 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 134 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 134; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 134, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta099 proteins include variants of SEQ ID NO: 134. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 134. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 134 while retaining at least one epitope of SEQ ID NO: 134. The first 30 N-terminal amino acids of SEQ ID NO: 134 can usefully be omitted. Other fragments omit one or more protein domains.

sta100

[0341] The 'sta100' antigen is annotated as 'superantigen-like protein'. In the NCTC 8325 strain sta100 is SAOUHSC_00391 and has amino acid sequence SEQ ID NO: 135 (GI:88194186).

[0342] Useful sta100 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 135 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 135; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 135, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta100 proteins include variants of SEQ ID NO: 135. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 135. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 135 while retaining at least one epitope of SEQ ID NO: 135. The first 30 N-terminal amino acids of SEQ ID NO: 135 can usefully be omitted. Other fragments omit one or more protein domains.

sta101

[0343] The 'sta101' antigen is annotated as 'superantigen-like protein 7'. In the NCTC 8325 strain sta101 is SAOUHSC_00392 and has amino acid sequence SEQ ID NO: 136 (GI:88194187). In the Newman strain it is nwmn_0394 (GI:151220606).

[0344] Useful sta101 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 136 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 136; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 136, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta101 proteins include variants of SEQ ID NO: 136. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 136. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 136 while retaining at least one epitope of SEQ ID NO: 136. The first 30 N-terminal amino acids of SEQ ID NO: 136 can usefully be omitted. Other fragments omit one or more protein domains.

sta102

[0345] The 'sta102' antigen is annotated as 'superantigen-like protein'. In the NCTC 8325 strain sta102 is SAOUHSC_00393 and has amino acid sequence SEQ ID NO: 137 (GI:88194188).

[0346] Useful sta102 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 137 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 137; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 137, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta102 proteins include variants of SEQ ID NO: 137. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 137. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 137 while retaining at least one epitope of SEQ ID NO: 137. The first 17 N-terminal amino acids of SEQ ID NO: 137 can usefully be omitted. Other fragments omit one or more protein domains.

sta103

[0347] The 'sta103' antigen is annotated as 'superantigen-like protein'. In the NCTC 8325 strain sta103 is SAOUHSC_00394 and has amino acid sequence SEQ ID NO: 138 (GI:88194189).

[0348] Useful sta103 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 138 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 138; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 138, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta103 proteins include variants of SEQ ID NO: 138. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 138. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 138 while retaining at least one epitope of SEQ ID NO: 138. The first 23 N-terminal amino acids of SEQ ID NO: 138 can usefully be omitted. Other fragments omit one or more protein domains.

sta104

[0349] The 'sta104' antigen is annotated as 'superantigen-like protein'. In the NCTC 8325 strain sta104 is SAOUHSC_00395 and has amino acid sequence SEQ ID NO: 139 (GI:88194190).

[0350] Useful sta104 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 139 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 139; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 139, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta104 proteins include variants of SEQ ID NO: 139. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 139. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 139 while retaining at least one epitope of SEQ ID NO: 139. Other fragments omit one or more protein domains.

sta105

[0351] The 'sta105' antigen is annotated as 'superantigen-like protein'. In the NCTC 8325 strain sta105 is SAOUHSC_00399 and has amino acid sequence SEQ ID NO: 140 (GI:88194194). In the Newman strain it is nwmn_0400 (GI:151220612).

[0352] Useful sta105 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 140 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 140; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 140, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta105 proteins include variants of SEQ ID NO: 140. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 140. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 140 while retaining at least one epitope of SEQ ID NO: 140. The first 30 N-terminal amino acids of SEQ ID NO: 140 can usefully be omitted.

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Other fragments omit one or more protein domains.

sta106

5 **[0353]** The 'sta106' antigen is annotated as 'hypothetical protein'. In the NCTC 8325 strain sta106 is SAOUHSC_01115 and has amino acid sequence SEQ ID NO: 141 (GI:88194861).

[0354] Useful sta106 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 141 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 141; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 141, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100 or more). These sta106 proteins include variants of SEQ ID NO: 141. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 141. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 141 while retaining at least one epitope of SEQ ID NO: 141. The first 16 N-terminal amino acids of SEQ ID NO: 141 can usefully be omitted. Other fragments omit one or more protein domains.

sta107

20 **[0355]** The 'sta107' antigen is annotated as 'hypothetical protein'. In the NCTC 8325 strain sta107 is SAOUHSC_00354 and has amino acid sequence SEQ ID NO: 177 (GI:88194153).

[0356] Useful sta107 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 177 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 177; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 177, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta107 proteins include variants of SEQ ID NO: 177. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 177. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 177 while retaining at least one epitope of SEQ ID NO: 177. The first 35 N-terminal amino acids of SEQ ID NO: 177 can usefully be omitted. Other fragments omit one or more protein domains.

sta108

35 **[0357]** The 'sta108' antigen is annotated as 'hypothetical protein'. In the NCTC 8325 strain sta108 is SAOUHSC_00717 and has amino acid sequence SEQ ID NO: 178 (GI:88194482).

[0358] Useful sta108 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 178 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 178; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 178, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100 or more). These sta108 proteins include variants of SEQ ID NO: 178. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 178. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 178 while retaining at least one epitope of SEQ ID NO: 178. The first 20 N-terminal amino acids of SEQ ID NO: 178 can usefully be omitted. Other fragments omit one or more protein domains.

sta109

50 **[0359]** The 'sta109' antigen is annotated as 'N-acetylmuramoyl-L-alanine amidase'. In the NCTC 8325 strain sta109 is SAOUHSC_02979 and has amino acid sequence SEQ ID NO: 179 (GI:88196599).

[0360] Useful sta109 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 179 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 179; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 179, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta109 proteins include variants of SEQ ID NO: 179. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 179. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or

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more amino acids (*e.g.* 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 179 while retaining at least one epitope of SEQ ID NO: 179. The first 27 N-terminal amino acids of SEQ ID NO: 179 can usefully be omitted. Other fragments omit one or more protein domains.

5 *sta110*

[0361] The 'sta110' antigen is annotated as 'hypothetical protein'. In the NCTC 8325 strain sta110 is SAOUHSC_01039 and has amino acid sequence SEQ ID NO: 180 (GI:88194791).

10 **[0362]** Useful sta110 antigens can elicit an antibody (*e.g.* when administered to a human) that recognises SEQ ID NO: 180 and/or may comprise an amino acid sequence: (a) having 50% or more identity (*e.g.* 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 180; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 180, wherein 'n' is 7 or more (*e.g.* 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These sta110 proteins include variants of SEQ ID NO: 180. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 180. Other preferred fragments lack
15 one or more amino acids (*e.g.* 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (*e.g.* 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 180 while retaining at least one epitope of SEQ ID NO: 180. The first 19 N-terminal amino acids of SEQ ID NO: 180 can usefully be omitted. Other fragments omit one or more protein domains.

20 *sta111*

[0363] The 'sta111' antigen is annotated as 'hypothetical protein'. In the NCTC 8325 strain sta111 is SAOUHSC_01005 and has amino acid sequence SEQ ID NO: 181 (GI:88194760).

25 **[0364]** Useful sta111 antigens can elicit an antibody (*e.g.* when administered to a human) that recognises SEQ ID NO: 181 and/or may comprise an amino acid sequence: (a) having 50% or more identity (*e.g.* 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 181; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 181, wherein 'n' is 7 or more (*e.g.* 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100 or more). These sta111 proteins include variants of SEQ ID NO: 181. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 181. Other preferred fragments lack one or
30 more amino acids (*e.g.* 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (*e.g.* 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 181 while retaining at least one epitope of SEQ ID NO: 181. The first 20 N-terminal amino acids of SEQ ID NO: 181 can usefully be omitted. Other fragments omit one or more protein domains.

35 *sta112*

[0365] The 'sta112' antigen is annotated as a putative 'ABC transporter, substrate-binding protein'. In the NCTC 8325 strain sta112 is SAOUHSC_00634 and has amino acid sequence SEQ ID NO: 182 (GI:88194402).

40 **[0366]** Useful sta112 antigens can elicit an antibody (*e.g.* when administered to a human) that recognises SEQ ID NO: 182 and/or may comprise an amino acid sequence: (a) having 50% or more identity (*e.g.* 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 182; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 182, wherein 'n' is 7 or more (*e.g.* 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta112 proteins include variants of SEQ ID NO: 182. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 182. Other preferred fragments
45 lack one or more amino acids (*e.g.* 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (*e.g.* 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 182 while retaining at least one epitope of SEQ ID NO: 182. The first 17 N-terminal amino acids of SEQ ID NO: 182 can usefully be omitted. Other fragments omit one or more protein domains.

[0367] A Sta112 antigen can usefully be included in a composition in combination with a Sta073 [74].

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sta113

[0368] The 'sta113' antigen is annotated as 'hypothetical protein'. In the NCTC 8325 strain sta113 is SAOUHSC_00728 and has amino acid sequence SEQ ID NO: 183 (GI:88194493).

55 **[0369]** Useful sta113 antigens can elicit an antibody (*e.g.* when administered to a human) that recognises SEQ ID NO: 183 and/or may comprise an amino acid sequence: (a) having 50% or more identity (*e.g.* 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 183; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 183, wherein 'n' is 7 or more (*e.g.* 8, 10,

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12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta113 proteins include variants of SEQ ID NO: 183. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 183. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 183 while retaining at least one epitope of SEQ ID NO: 183. The first 173 N-terminal amino acids of SEQ ID NO: 183 can usefully be omitted. Other fragments omit one or more protein domains.

sta114

[0370] The 'sta114' antigen is annotated as 'hypothetical protein'. In the NCTC 8325 strain sta114 is SAOUHSC_00810 and has amino acid sequence SEQ ID NO: 184 (GI:88194570).

[0371] Useful sta114 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 184 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 184; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 184, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150 or more). These sta114 proteins include variants of SEQ ID NO: 184. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 184. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 184 while retaining at least one epitope of SEQ ID NO: 184. Other fragments omit one or more protein domains.

sta115

[0372] The 'sta115' antigen is annotated as 'hypothetical protein'. In the NCTC 8325 strain sta115 is SAOUHSC_00817 and has amino acid sequence SEQ ID NO: 185 (GI:88194576).

[0373] Useful sta115 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 185 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 185; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 185, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150 or more). These sta115 proteins include variants of SEQ ID NO: 185. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 185. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 185 while retaining at least one epitope of SEQ ID NO: 185. The first 18 N-terminal amino acids of SEQ ID NO: 185 can usefully be omitted. Other fragments omit one or more protein domains.

sta116

[0374] The 'sta116' antigen is annotated as 'formyl peptide receptor-like 1 inhibitory protein'. In the NCTC 8325 strain sta116 is SAOUHSC_01112 and has amino acid sequence SEQ ID NO: 186 (GI:88194858).

[0375] Useful sta116 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 186 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 186; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 186, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100 or more). These sta116 proteins include variants of SEQ ID NO: 186. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 186. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 186 while retaining at least one epitope of SEQ ID NO: 186. The first 20 N-terminal amino acids of SEQ ID NO: 186 can usefully be omitted. Other fragments omit one or more protein domains.

sta117

[0376] The 'sta117' antigen is annotated as 'truncated beta-hemolysin'. In the NCTC 8325 strain sta117 is SAOUHSC_02240 and has amino acid sequence SEQ ID NO: 187 (GI:88195913).

[0377] Useful sta117 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 187 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 187; and/or (b)

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comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 187, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta117 proteins include variants of SEQ ID NO: 187. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 187. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 187 while retaining at least one epitope of SEQ ID NO: 187. Other fragments omit one or more protein domains.

sta118

[0378] The 'sta118' antigen is annotated as 'cell division protein FtsZ'. In the NCTC 8325 strain sta118 is SAOUHSC_01150 and has amino acid sequence SEQ ID NO: 188 (GI:88194892).

[0379] Useful sta118 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 188 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 188; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 188, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These sta118 proteins include variants of SEQ ID NO: 188. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 188. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 188 while retaining at least one epitope of SEQ ID NO: 188. Other fragments omit one or more protein domains.

sta119

[0380] The 'sta119' antigen is annotated as 'thioredoxin'. In the NCTC 8325 strain sta119 is SAOUHSC_01100 and has amino acid sequence SEQ ID NO: 200 (GI:88194846).

[0381] Useful sta119 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 200 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 200; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 200, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100 or more). These sta119 proteins include variants of SEQ ID NO: 200. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 200. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 200 while retaining at least one epitope of SEQ ID NO: 200. Other fragments omit one or more protein domains.

sta120

[0382] The 'sta120' antigen is annotated as 'alkyl hydroperoxide reductase subunit C'. In the NCTC 8325 strain sta120 is SAOUHSC_00365 and has amino acid sequence SEQ ID NO: 201 (GI:88194163).

[0383] Useful sta120 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 201 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 201; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 201, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150 or more). These sta120 proteins include variants of SEQ ID NO: 201. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 201. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 201 while retaining at least one epitope of SEQ ID NO: 201. Other fragments omit one or more protein domains.

NW_6

[0384] The 'NW_6' antigen is annotated as 'secreted von Willebrand factor-binding protein precursor'. In the Newman strain NW_6 is NWMN_0757 and has amino acid sequence SEQ ID NO: 142 (GI: 151220969).

[0385] Useful NW_6 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 142 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 142; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 142, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These NW_6 proteins include variants

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of SEQ ID NO: 142. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 142. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 142 while retaining at least one epitope of SEQ ID NO: 142. The first 13 N-terminal amino acids of SEQ ID NO: 142 can usefully be omitted. Other fragments omit one or more protein domains.

NW_9

[0386] The 'NW_9' antigen is annotated as 'lipoprotein'. In the Newman strain NW_9 is NWMN_0958 and has amino acid sequence SEQ ID NO: 143 (GI: 151221170).

[0387] Useful NW_9 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 143 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 143; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 143, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200 or more). These NW_9 proteins include variants of SEQ ID NO: 143. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 143. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 143 while retaining at least one epitope of SEQ ID NO: 143. The first 19 N-terminal amino acids of SEQ ID NO: 143 can usefully be omitted. Other fragments omit one or more protein domains.

NW_10

[0388] The 'NW_10' antigen is annotated as 'fibrinogen binding-related protein'. In the Newman strain NW_10 is NWMN_1066 and has amino acid sequence SEQ ID NO: 144 (GI: 151221278).

[0389] Useful NW_10 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 144 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 144; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 144, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100 or more). These NW_10 proteins include variants of SEQ ID NO: 144. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 144. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 144 while retaining at least one epitope of SEQ ID NO: 144. The first 20 N-terminal amino acids of SEQ ID NO: 144 can usefully be omitted. Other fragments omit one or more protein domains.

NW_7

[0390] The 'NW_7' antigen is annotated as 'staphylococcal complement inhibitor SCIN'. In the Newman strain NW_7 is NWMN_1876 and has amino acid sequence SEQ ID NO: 145 (GI:151222088).

[0391] Useful NW_7 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 145 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 145; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 145, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100 or more). These NW_7 proteins include variants of SEQ ID NO: 145. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 145. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 145 while retaining at least one epitope of SEQ ID NO: 145. The first 17 N-terminal amino acids of SEQ ID NO: 145 can usefully be omitted. Other fragments omit one or more protein domains.

NW_8

[0392] The 'NW_8' antigen is annotated as 'chemotaxis-inhibiting protein CHIPS'. In the Newman strain NW_8 is NWMN_1877 and has amino acid sequence SEQ ID NO: 146 (GI:151222089).

[0393] Useful NW_8 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 146 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 146; and/or (b)

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comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 146, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100 or more). These NW_8 proteins include variants of SEQ ID NO: 146. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 146. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 146 while retaining at least one epitope of SEQ ID NO: 146. The first 19 N-terminal amino acids of SEQ ID NO: 146 can usefully be omitted. Other fragments omit one or more protein domains.

NW_2

[0394] The 'NW_2' antigen is annotated as 'enterotoxin type A precursor'. In the Newman strain NW_2 is NWMN_1883 and has amino acid sequence SEQ ID NO: 147 (GI: 151222095).

[0395] Useful NW_2 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 147 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 147; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 147, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These NW_2 proteins include variants of SEQ ID NO: 147. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 147. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 147 while retaining at least one epitope of SEQ ID NO: 147. The first 16 N-terminal amino acids of SEQ ID NO: 147 can usefully be omitted. Other fragments omit one or more protein domains.

NW_1

[0396] The 'NW_1' antigen is annotated as 'lipoprotein'. In the Newman strain NW_1 is NWMN_1924 and has amino acid sequence SEQ ID NO: 148 (GI: 151222136).

[0397] Useful NW_1 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 148 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 148; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 148, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150 or more). These NW_1 proteins include variants of SEQ ID NO: 148. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 148. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 148 while retaining at least one epitope of SEQ ID NO: 148. The first 17 N-terminal amino acids of SEQ ID NO: 148 can usefully be omitted. Other fragments omit one or more protein domains.

NW_5

[0398] The 'NW_5' antigen is annotated as 'cell wall surface anchor family protein'. In the Newman strain NW_5 is NWMN_2392 and has amino acid sequence SEQ ID NO: 149 (GI:151222604).

[0399] Useful NW_5 antigens can elicit an antibody (e.g. when administered to a human) that recognises SEQ ID NO: 149 and/or may comprise an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 149; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 149, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more). These NW_5 proteins include variants of SEQ ID NO: 149. Preferred fragments of (b) comprise an epitope from SEQ ID NO: 149. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 149 while retaining at least one epitope of SEQ ID NO: 149. The first 52 N-terminal amino acids of SEQ ID NO: 149 can usefully be omitted. Other fragments omit one or more protein domains.

Hybrid polypeptides

[0400] Antigens used in the invention may be present in the composition as individual separate polypeptides. Where more than one antigen is used, however, they do not have to be present as separate polypeptides. Instead, at least two (e.g. 2, 3, 4, 5, or more) antigens can be expressed as a single polypeptide chain (a 'hybrid' polypeptide). Hybrid

polypeptides offer two main advantages: first, a polypeptide that may be unstable or poorly expressed on its own can be assisted by adding a suitable hybrid partner that overcomes the problem; second, commercial manufacture is simplified as only one expression and purification need be employed in order to produce two polypeptides which are both antigenically useful.

[0401] The hybrid polypeptide may comprise two or more polypeptide sequences from the first antigen group. The hybrid polypeptide may comprise one or more polypeptide sequences from the first antigen group and one or more polypeptide sequences from the second antigen group. Moreover, the hybrid polypeptide may comprise two or more polypeptide sequences from each of the antigens listed above, or two or more variants of the same antigen in the cases in which the sequence has partial variability across strains.

[0402] Hybrids consisting of amino acid sequences from two, three, four, five, six, seven, eight, nine, or ten antigens are useful. In particular, hybrids consisting of amino acid sequences from two, three, four, or five antigens are preferred, such as two or three antigens.

[0403] Different hybrid polypeptides may be mixed together in a single formulation. Hybrids may be combined with non-hybrid antigens selected from the first, second or third antigen groups. Within such combinations, an antigen may be present in more than one hybrid polypeptide and/or as a non-hybrid polypeptide. It is preferred, however, that an antigen is present either as a hybrid or as a non-hybrid, but not as both.

[0404] The hybrid polypeptides can also be combined with conjugates or non-*S.aureus* antigens as described above.

[0405] Hybrid polypeptides can be represented by the formula $\text{NH}_2\text{-A}\{-\text{X-L}\}_n\text{-B-COOH}$, wherein: X is an amino acid sequence of a *S.aureus* antigen, as described above; L is an optional linker amino acid sequence; A is an optional N-terminal amino acid sequence; B is an optional C-terminal amino acid sequence; n is an integer of 2 or more (e.g. 2, 3, 4, 5, 6, etc.). Usually n is 2 or 3.

[0406] If a -X- moiety has a leader peptide sequence in its wild-type form, this may be included or omitted in the hybrid protein. In some embodiments, the leader peptides will be deleted except for that of the -X- moiety located at the N-terminus of the hybrid protein i.e. the leader peptide of X_1 will be retained, but the leader peptides of $X_2 \dots X_n$ will be omitted. This is equivalent to deleting all leader peptides and using the leader peptide of X_1 as moiety -A-.

[0407] For each n instances of {-X-L-}, linker amino acid sequence -L- may be present or absent. For instance, when n=2 the hybrid may be $\text{NH}_2\text{-X}_1\text{-L}_1\text{-X}_2\text{-L}_2\text{-COOH}$, $\text{NH}_2\text{-X}_1\text{-X}_2\text{-COOH}$, $\text{NH}_2\text{-X}_1\text{-L}_1\text{-X}_2\text{-COOH}$, $\text{NH}_2\text{-X}_1\text{-X}_2\text{-L}_2\text{-COOH}$, etc. Linker amino acid sequence(s) -L- will typically be short (e.g. 20 or fewer amino acids i.e. 20, 19, 18, 17, 16, 15, 14, 13, 12, 11, 10, 9, 8, 7, 6, 5, 4, 3, 2, 1). Examples comprise short peptide sequences which facilitate cloning, poly-glycine linkers (i.e. comprising Gly_n where n = 2, 3, 4, 5, 6, 7, 8, 9, 10 or more), and histidine tags (i.e. His_n where n = 3, 4, 5, 6, 7, 8, 9, 10 or more). Other suitable linker amino acid sequences will be apparent to those skilled in the art. A useful linker is GSGGGG (SEQ ID NO: 171) or GSGSGGGG (SEQ ID NO: 172), with the Gly-Ser dipeptide being formed from a *Bam*HI restriction site (or two of them, to form the SEQ ID NO: 230 tetrapeptide), thus aiding cloning and manipulation, and the $(\text{Gly})_4$ tetrapeptide (SEQ ID NO: 227) being a typical poly-glycine linker. Other suitable linkers, particularly for use as the final L_n are ASGGGS (SEQ ID NO: 173 e.g. encoded by SEQ ID NO: 174) or a Leu-Glu dipeptide.

[0408] -A- is an optional N-terminal amino acid sequence. This will typically be short (e.g. 40 or fewer amino acids i.e. 40, 39, 38, 37, 36, 35, 34, 33, 32, 31, 30, 29, 28, 27, 26, 25, 24, 23, 22, 21, 20, 19, 18, 17, 16, 15, 14, 13, 12, 11, 10, 9, 8, 7, 6, 5, 4, 3, 2, 1). Examples include leader sequences to direct protein trafficking, or short peptide sequences which facilitate cloning or purification (e.g. histidine tags i.e. His_n where n = 3, 4, 5, 6, 7, 8, 9, 10 or more). Other suitable N-terminal amino acid sequences will be apparent to those skilled in the art. If X_1 lacks its own N-terminus methionine, -A- is preferably an oligopeptide (e.g. with 1, 2, 3, 4, 5, 6, 7 or 8 amino acids) which provides a N-terminus methionine e.g. Met-Ala-Ser, or a single Met residue.

[0409] -B- is an optional C-terminal amino acid sequence. This will typically be short (e.g. 40 or fewer amino acids i.e. 39, 38, 37, 36, 35, 34, 33, 32, 31, 30, 29, 28, 27, 26, 25, 24, 23, 22, 21, 20, 19, 18, 17, 16, 15, 14, 13, 12, 11, 10, 9, 8, 7, 6, 5, 4, 3, 2, 1). Examples include sequences to direct protein trafficking, short peptide sequences which facilitate cloning or purification (e.g. comprising histidine tags i.e. His_n where n = 3, 4, 5, 6, 7, 8, 9, 10 or more, such as SEQ ID NO: 226), or sequences which enhance protein stability. Other suitable C-terminal amino acid sequences will be apparent to those skilled in the art.

[0410] One hybrid polypeptide of the invention may include both EsxA and EsxB antigens. These may be in either order, N- to C- terminus. SEQ ID NOs: 151 ('EsxAB'; encoded by SEQ ID NO: 169) and 152 ('EsxBA') are examples of such hybrids, both having hexapeptide linkers ASGGGS (SEQ ID NO: 173). Another 'EsxAB' hybrid comprises SEQ ID NO: 241, which may be provided with a N-terminus methionine (e.g. SEQ ID NO: 250).

[0411] Another hybrid polypeptide of the invention may include both SdrD and SdrE antigens. These may be in either order, N- to C- terminus. SEQ ID NO: 168 ('SdrED') is an example of such a hybrid, having a hexapeptide linker ASGGGS (SEQ ID NO: 173).

[0412] Another hybrid polypeptide of the invention may include both ClfB and SdrD antigens. These may be in either order, N- to C- terminus. SEQ ID NO: 202 ('ClfB-SdrD') is an example of such a hybrid, having a hexapeptide linker ASGGGS (SEQ ID NO: 173). SEQ ID NO: 203 ('SdrD-ClfB') is another example of such a hybrid, having a hexapeptide

linker ASGGGS (SEQ ID NO: 173). SEQ ID NO: 211 ('ClfB-N3-sdrD-N3') is another example of such a hybrid, where the N3 fragments of ClfB and SdrD are joined by hexapeptide linker ASGGGS (SEQ ID NO: 173).

[0413] Another hybrid polypeptide of the invention may include both LsdA and EsxA antigens. These may be in either order, N- to C- terminus. SEQ ID NO: 204 ('LsdA-EsxA') is an example of such a hybrid, having a hexapeptide linker ASGGGS (SEQ ID NO: 173). SEQ ID NO: 209 ('isdA40-184-esxA') is another example of such a hybrid, in which LsdA₄₀₋₁₈₄ is joined to EsxA via linker ASGGGS (SEQ ID NO: 173).

[0414] Another hybrid polypeptide of the invention may include both LsdA and sta006 antigens. These may be in either order, N- to C- terminus. SEQ ID NO: 221 ('isdA40-184-sta006') is an example of such a hybrid, in which LsdA₄₀₋₁₈₄ is joined to Sta006 via hexapeptide linker ASGGGS (SEQ ID NO: 173).

[0415] Another hybrid polypeptide of the invention may include both Hla and sta006 antigens. These may be in either order, N- to C- terminus. SEQ ID NO: 222 ('HlaH35L-sta006') is an example of such a hybrid, in which a H35L mutant of H1a is joined to Sta006 via hexapeptide linker ASGGGS (SEQ ID NO: 173).

[0416] Another hybrid polypeptide of the invention may include both Hla and Emp antigens. These may be in either order, N- to C- terminus. SEQ ID NO: 205 ('HlaH35L-Emp') is an example of such a hybrid, in which a H35L mutant Hla is joined to Emp via linker ASGGGS (SEQ ID NO: 173). SEQ ID NO: 206 ('Hla27-76-Emp') is another example of such a hybrid, in which a Hla fragment is joined to Emp via linker ASGGGS (SEQ ID NO: 173); SEQ ID NO: 207 is a H35L mutant of SEQ ID NO: 206. SEQ ID NO: 208 ('HlaPSGS-Emp') is another example of such a hybrid, in which a Hla mutant is joined to Emp via linker ASGGGS (SEQ ID NO: 173).

[0417] Another hybrid polypeptide of the invention may include LsdA and EsxA and EsxB antigens. These may be in any order, N- to C- terminus. SEQ ID NO: 210 ('isdA40-184-esxAB') is an example of such a triple hybrid, in which LsdA₄₀₋₁₈₄ is joined to EsxAB via linker ASGGGS (SEQ ID NO: 173). The EsxAB already includes the same linker, so SEQ ID NO: 210 includes two of these linkers. SEQ ID NO: 212 ('LsdA-esxAB') is another example of such a triple hybrid, in which LsdA is joined to EsxAB via linker ASGGGS (SEQ ID NO: 173).

[0418] Another hybrid polypeptide of the invention may include Hla and EsxA and EsxB antigens. These may be in any order, N- to C- terminus. SEQ ID NO: 220 ('HlaH35L-esxAB') is an example of such a triple hybrid, in which a H35L mutant of H1a is joined to EsxAB via linker ASGGGS (SEQ ID NO: 173). The EsxAB already includes the same linker, so SEQ ID NO: 220 includes two of these linkers. Another example of a hybrid polypeptide including Hla and EsxA and EsxB antigens is SEQ ID NO: 237 ('HlaH35L-esxAB' as used in the examples), in which a H35L mutant of H1a is joined to EsxA via linker APTARG (SEQ ID NO: 239) to replace its N-terminus, then to EsxB via linker ASGGGS (SEQ ID NO: 173) to replace its N-terminus. This hybrid can be provided with a suitable N-terminal sequence such as SEQ ID NO: 240.

[0419] Another hybrid polypeptide of the invention may include sta006 and EsxA and EsxB antigens. These may be in any order, N- to C- terminus. SEQ ID NO: 223 ('sta006-esxAB') is an example of such a triple hybrid, in which sta006 is joined to EsxAB via linker ASGGGS (SEQ ID NO: 173). The EsxAB already includes the same linker, so SEQ ID NO: 223 includes two of these linkers. Another example of a hybrid polypeptide including sta006 and EsxA and EsxB antigens is SEQ ID NO: 238 ('sta006-esxAB' as used in the examples), in which a sta006 is joined to EsxA via linker APTARG (SEQ ID NO: 239) to replace its N-terminus, then to EsxB via linker ASGGGS (SEQ ID NO: 173) to replace its N-terminus. This hybrid can be provided with a suitable N-terminal sequence such as SEQ ID NO: 240.

[0420] Usefully, these hybrid polypeptides can elicit an antibody (e.g. when administered to a human) that recognise each of the wild-type staphylococcal proteins (e.g. as shown in the sequence listing) represented in the hybrid e.g. which recognise both wild-type EsxA and wild-type EsxB, or which recognise both wild-type SdrD and wild-type SdrE, or which recognise both wild-type SdrD and wild-type ClfB, or which recognise both wild-type LsdA and wild-type EsxA, or which recognise both wild-type LsdA and wild-type sta006, or which recognise both wild-type Hla and wild-type sta006, or which recognise both wild-type Hla and wild-type Emp, or which recognise wild-type LsdA and wild-type EsxA and wild-type EsxB, or which recognise wild-type Hla and wild-type EsxA and wild-type EsxB, or which recognise wild-type sta006 and wild-type EsxA and wild-type EsxB.

Polypeptides used with the invention

[0421] Polypeptides used with the invention can take various forms (e.g. native, fusions, glycosylated, non-glycosylated, lipidated, non-lipidated, phosphorylated, non-phosphorylated, myristoylated, non-myristoylated, monomeric, multimeric, particulate, denatured, etc.).

[0422] Polypeptides used with the invention can be prepared by various means (e.g. recombinant expression, purification from cell culture, chemical synthesis, etc.). Recombinantly-expressed proteins are preferred, particularly for hybrid polypeptides.

[0423] Polypeptides used with the invention are preferably provided in purified or substantially purified form i.e. substantially free from other polypeptides (e.g. free from naturally-occurring polypeptides), particularly from other staphylococcal or host cell polypeptides, and are generally at least about 50% pure (by weight), and usually at least about 90% pure i.e. less than about 50%, and more preferably less than about 10% (e.g. 5%) of a composition is made up of other

expressed polypeptides. Thus the antigens in the compositions are separated from the whole organism with which the molecule is expressed.

[0424] Polypeptides used with the invention are preferably staphylococcal polypeptides.

[0425] The term "polypeptide" refers to amino acid polymers of any length. The polymer may be linear or branched, it may comprise modified amino acids, and it may be interrupted by non-amino acids. The terms also encompass an amino acid polymer that has been modified naturally or by intervention; for example, disulfide bond formation, glycosylation, lipidation, acetylation, phosphorylation, or any other manipulation or modification, such as conjugation with a labeling component. Also included are, for example, polypeptides containing one or more analogs of an amino acid (including, for example, unnatural amino acids, *etc.*), as well as other modifications known in the art. Polypeptides can occur as single chains or associated chains.

[0426] The invention provides polypeptides comprising a sequence -P-Q- or -Q-P-, wherein: -P- is an amino acid sequence as defined above and -Q- is not a sequence as defined above *i. e.* the invention provides fusion proteins. Where the N-terminus codon of -P- is not ATG, but this codon is not present at the N-terminus of a polypeptide, it will be translated as the standard amino acid for that codon rather than as a Met. Where this codon is at the N-terminus of a polypeptide, however, it will be translated as Met. Examples of -Q- moieties include, but are not limited to, histidine tags (*i.e.* His_n, where *n* = 3, 4, 5, 6, 7, 8, 9, 10 or more), maltose-binding protein, or glutathione-S-transferase (GST).

[0427] The invention also provides a process for producing a polypeptide of the invention, comprising the step of culturing a host cell transformed with nucleic acid of the invention under conditions which induce polypeptide expression.

[0428] Although expression of the polypeptides of the invention may take place in a *Staphylococcus*, the invention will usually use a heterologous host for expression (recombinant expression). The heterologous host may be prokaryotic (*e.g.* a bacterium) or eukaryotic. It may be *E.coli*, but other suitable hosts include *Bacillus subtilis*, *Vibrio cholerae*, *Salmonella typhi*, *Salmonella typhimurium*, *Neisseria lactamica*, *Neisseria cinerea*, *Mycobacteria* (*e.g.* *M.tuberculosis*), yeasts, *etc.* Compared to the wild-type *S.aureus* genes encoding polypeptides of the invention, it is helpful to change codons to optimise expression efficiency in such hosts without affecting the encoded amino acids.

[0429] The invention provides a process for producing a polypeptide of the invention, comprising the step of synthesising at least part of the polypeptide by chemical means.

Nucleic acids

[0430] The invention also provides nucleic acid encoding polypeptides and hybrid polypeptides of the invention. It also provides nucleic acid comprising a nucleotide sequence that encodes one or more polypeptides or hybrid polypeptides of the invention.

[0431] The invention also provides nucleic acid comprising nucleotide sequences having sequence identity to such nucleotide sequences. Identity between sequences is preferably determined by the Smith-Waterman homology search algorithm as described above. Such nucleic acids include those using alternative codons to encode the same amino acid.

[0432] The invention also provides nucleic acid which can hybridize to these nucleic acids. Hybridization reactions can be performed under conditions of different "stringency". Conditions that increase stringency of a hybridization reaction of widely known and published in the art (*e.g.* page 7.52 of reference 276). Examples of relevant conditions include (in order of increasing stringency): incubation temperatures of 25°C, 37°C, 50°C, 55°C and 68°C; buffer concentrations of 10 x SSC, 6 x SSC, 1 x SSC, 0.1 x SSC (where SSC is 0.15 M NaCl and 15 mM citrate buffer) and their equivalents using other buffer systems; formamide concentrations of 0%, 25%, 50%, and 75%; incubation times from 5 minutes to 24 hours; 1, 2, or more washing steps; wash incubation times of 1, 2, or 15 minutes; and wash solutions of 6 x SSC, 1 x SSC, 0.1 x SSC, or de-ionized water. Hybridization techniques and their optimization are well known in the art (*e.g.* see refs 75, 76, 276, 278, *etc.*).

[0433] In some embodiments, nucleic acid of the invention hybridizes to a target under low stringency conditions; in other embodiments it hybridizes under intermediate stringency conditions; in preferred embodiments, it hybridizes under high stringency conditions. An exemplary set of low stringency hybridization conditions is 50°C and 10 x SSC. An exemplary set of intermediate stringency hybridization conditions is 55°C and 1 x SSC. An exemplary set of high stringency hybridization conditions is 68°C and 0.1 x SSC.

[0434] The invention includes nucleic acid comprising sequences complementary to these sequences (*e.g.* for antisense or probing, or for use as primers).

[0435] Nucleic acids of the invention can be used in hybridisation reactions (*e.g.* Northern or Southern blots, or in nucleic acid microarrays or 'gene chips') and amplification reactions (*e.g.* PCR, SDA, SSSR, LCR, TMA, NASBA, *etc.*) and other nucleic acid techniques.

[0436] Nucleic acid according to the invention can take various forms (*e.g.* single-stranded, double-stranded, vectors, primers, probes, labelled *etc.*). Nucleic acids of the invention may be circular or branched, but will generally be linear. Unless otherwise specified or required, any embodiment of the invention that utilizes a nucleic acid may utilize both the double-stranded form and each of two complementary single-stranded forms which make up the double-stranded form.

Primers and probes are generally single-stranded, as are antisense nucleic acids.

[0437] Nucleic acids of the invention are preferably provided in purified or substantially purified form *i. e.* substantially free from other nucleic acids (*e.g.* free from naturally-occurring nucleic acids), particularly from other staphylococcal or host cell nucleic acids, generally being at least about 50% pure (by weight), and usually at least about 90% pure. Nucleic acids of the invention are preferably staphylococcal nucleic acids.

[0438] Nucleic acids of the invention may be prepared in many ways *e.g.* by chemical synthesis (*e.g.* phosphoramidite synthesis of DNA) in whole or in part, by digesting longer nucleic acids using nucleases (*e.g.* restriction enzymes), by joining shorter nucleic acids or nucleotides (*e.g.* using ligases or polymerases), from genomic or cDNA libraries, *etc.*

[0439] Nucleic acid of the invention may be attached to a solid support (*e.g.* a bead, plate, filter, film, slide, microarray support, resin, *etc.*). Nucleic acid of the invention may be labelled *e.g.* with a radioactive or fluorescent label, or a biotin label. This is particularly useful where the nucleic acid is to be used in detection techniques *e.g.* where the nucleic acid is a primer or as a probe.

[0440] The term "nucleic acid" includes in general means a polymeric form of nucleotides of any length, which contain deoxyribonucleotides, ribonucleotides, and/or their analogs. It includes DNA, RNA, DNA/RNA hybrids. It also includes DNA or RNA analogs, such as those containing modified backbones (*e.g.* peptide nucleic acids (PNAs) or phosphorothioates) or modified bases. Thus the invention includes mRNA, tRNA, rRNA, ribozymes, DNA, cDNA, recombinant nucleic acids, branched nucleic acids, plasmids, vectors, probes, primers, *etc.* Where nucleic acid of the invention takes the form of RNA, it may or may not have a 5' cap.

[0441] Nucleic acids of the invention may be part of a vector *i.e.* part of a nucleic acid construct designed for transduction/transfection of one or more cell types. Vectors may be, for example, "cloning vectors" which are designed for isolation, propagation and replication of inserted nucleotides, "expression vectors" which are designed for expression of a nucleotide sequence in a host cell, "viral vectors" which is designed to result in the production of a recombinant virus or virus-like particle, or "shuttle vectors", which comprise the attributes of more than one type of vector. Preferred vectors are plasmids. A "host cell" includes an individual cell or cell culture which can be or has been a recipient of exogenous nucleic acid. Host cells include progeny of a single host cell, and the progeny may not necessarily be completely identical (in morphology or in total DNA complement) to the original parent cell due to natural, accidental, or deliberate mutation and/or change. Host cells include cells transfected or infected *in vivo* or *in vitro* with nucleic acid of the invention.

[0442] Where a nucleic acid is DNA, it will be appreciated that "U" in a RNA sequence will be replaced by "T" in the DNA. Similarly, where a nucleic acid is RNA, it will be appreciated that "T" in a DNA sequence will be replaced by "U" in the RNA.

[0443] The term "complement" or "complementary" when used in relation to nucleic acids refers to Watson-Crick base pairing. Thus the complement of C is G, the complement of G is C, the complement of A is T (or U), and the complement of T (or U) is A. It is also possible to use bases such as I (the purine inosine) *e.g.* to complement pyrimidines (C or T).

[0444] Nucleic acids of the invention can be used, for example: to produce polypeptides; as hybridization probes for the detection of nucleic acid in biological samples; to generate additional copies of the nucleic acids; to generate ribozymes or antisense oligonucleotides; as single-stranded DNA primers or probes; or as triple-strand forming oligonucleotides.

[0445] The invention provides a process for producing nucleic acid of the invention, wherein the nucleic acid is synthesised in part or in whole using chemical means.

[0446] The invention provides vectors comprising nucleotide sequences of the invention (*e.g.* cloning or expression vectors) and host cells transformed with such vectors.

[0447] Nucleic acid amplification according to the invention may be quantitative and/or real-time.

[0448] For certain embodiments of the invention, nucleic acids are preferably at least 7 nucleotides in length (*e.g.* 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 20, 21, 22, 23, 24, 25, 26, 27, 28, 29, 30, 31, 32, 33, 34, 35, 36, 37, 38, 39, 40, 45, 50, 55, 60, 65, 70, 75, 80, 90, 100, 110, 120, 130, 140, 150, 160, 170, 180, 190, 200, 225, 250, 275, 300 nucleotides or longer).

[0449] For certain embodiments of the invention, nucleic acids are preferably at most 500 nucleotides in length (*e.g.* 450, 400, 350, 300, 250, 200, 150, 140, 130, 120, 110, 100, 90, 80, 75, 70, 65, 60, 55, 50, 45, 40, 39, 38, 37, 36, 35, 34, 33, 32, 31, 30, 29, 28, 27, 26, 25, 24, 23, 22, 21, 20, 19, 18, 17, 16, 15 nucleotides or shorter).

[0450] Primers and probes of the invention, and other nucleic acids used for hybridization, are preferably between 10 and 30 nucleotides in length (*e.g.* 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 20, 21, 22, 23, 24, 25, 26, 27, 28, 29, or 30 nucleotides).

Strains and variants

[0451] Antigens are defined above by reference to existing nomenclature (*e.g.* "ClfA"), to "sta" numbers or to "NW_" numbers. Table 1 herein relates these three naming/numbering systems to existing

[0452] SAOUHSC numbering and/or NWMN numbering. SAOUHSC numbering refers to the genome of *S.aureus* strain NCTC 8325 (sequenced by Oklahoma University Health Sciences Center and disclosed in GenBank as

CP000253.1; GI:87201381), and individual SAOUHSC numbers are given as "locus_tag" entries in the genome sequence's "features" section. Similarly, NWMN numbering refers to the genome of *S.aureus* strain Newman (isolated in 1952 from a human infection, and having robust virulence phenotype) disclosed in GenBank as AP009351.1 (GI: 150373012) and individual NWMN numbers are given as "locus_tag" entries in the genome sequence's "features" section. Functional annotations for each antigen are also given in the databases.

[0453] Table 1 also includes the GI number for each antigen of the invention. Thus an exemplary amino acid and nucleotide sequence for any of these antigens can easily be found in public sequence databases from the NCTC 8325 and/or Newman strain, but the invention is not limited to sequences from the NCTC 8325 and Newman strains. Genome sequences of several other strains of *S.aureus* are available, including those of MRSA strains N315 and Mu50 [77], MW2, N315, COL, MRSA252, MSSA476, RF122, USA300 (very virulent), JH1 and JH9. Standard search and alignment techniques can be used to identify in any of these (or other) further genome sequences the homolog of any particular sequence from the Newman or NCTC 8325 strain. Moreover, the available sequences from the Newman and NCTC 8325 strains can be used to design primers for amplification of homologous sequences from other strains. Thus the invention is not limited to these two strains, but rather encompasses such variants and homologs from other strains of *S.aureus*, as well as non-natural variants. In general, suitable variants of a particular SEQ ID NO include its allelic variants, its polymorphic forms, its homologs, its orthologs, its paralogs, its mutants, etc.

[0454] Thus, for instance, polypeptides used with the invention may, compared to the SEQ ID NO herein, include one or more (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, etc.) amino acid substitutions, such as conservative substitutions (*i. e.* substitutions of one amino acid with another which has a related side chain). Genetically-encoded amino acids are generally divided into four families: (1) acidic *i.e.* aspartate, glutamate; (2) basic *i.e.* lysine, arginine, histidine; (3) non-polar *i.e.* alanine, valine, leucine, isoleucine, proline, phenylalanine, methionine, tryptophan; and (4) uncharged polar *i.e.* glycine, asparagine, glutamine, cysteine, serine, threonine, tyrosine. Phenylalanine, tryptophan, and tyrosine are sometimes classified jointly as aromatic amino acids. In general, substitution of single amino acids within these families does not have a major effect on the biological activity. The polypeptides may also include one or more (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, etc.) single amino acid deletions relative to the SEQ ID NO sequences. The polypeptides may also include one or more (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, etc.) insertions (e.g. each of 1, 2, 3, 4 or 5 amino acids) relative to the SEQ ID NO sequences.

[0455] Similarly, a polypeptide used with the invention may comprise an amino acid sequence that:

is identical (*i. e.* 100% identical) to a sequence disclosed in the sequence listing;

shares sequence identity (e.g. 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) with a sequence disclosed in the sequence listing;

has 1, 2, 3, 4, 5, 6, 7, 8, 9 or 10 (or more) single amino acid alterations (deletions, insertions, substitutions), which may be at separate locations or may be contiguous, as compared to the sequences of (a) or (b); and

when aligned with a particular sequence from the sequence listing using a pairwise alignment algorithm, each moving window of x amino acids from N-terminus to C-terminus (such that for an alignment that extends to p amino acids, where $p > x$, there are $p-x+1$ such windows) has at least $x \cdot y$ identical aligned amino acids, where: x is selected from 20, 25, 30, 35, 40, 45, 50, 60, 70, 80, 90, 100, 150, 200; y is selected from 0.50, 0.60, 0.70, 0.75, 0.80, 0.85, 0.90, 0.91, 0.92, 0.93, 0.94, 0.95, 0.96, 0.97, 0.98, 0.99; and if $x \cdot y$ is not an integer then it is rounded up to the nearest integer. The preferred pairwise alignment algorithm is the Needleman-Wunsch global alignment algorithm [78], using default parameters (e.g. with Gap opening penalty = 10.0, and with Gap extension penalty = 0.5, using the EBLOSUM62 scoring matrix). This algorithm is conveniently implemented in the *needle* tool in the EMBOSS package [79].

[0456] Where hybrid polypeptides are used, the individual antigens within the hybrid (*i.e.* individual -X-moieties) may be from one or more strains. Where $n=2$, for instance, X_2 may be from the same strain as X_1 or from a different strain. Where $n=3$, the strains might be (i) $X_1=X_2=X_3$ (ii) $X_1=X_2 \neq X_3$ (iii) $X_1 \neq X_2 = X_3$ (iv) $X_1 \neq X_2 \neq X_3$ or (v) $X_1 = X_3 \neq X_2$, etc.

[0457] Within group (c), deletions or substitutions may be at the N-terminus and/or C-terminus, or may be between the two termini. Thus a truncation is an example of a deletion. Truncations may involve deletion of up to 40 (or more) amino acids at the N-terminus and/or C-terminus. N-terminus truncation can remove leader peptides e.g. to facilitate recombinant expression in a heterologous host. C-terminus truncation can remove anchor sequences e.g. to facilitate recombinant expression in a heterologous host.

[0458] In general, when an antigen comprises a sequence that is not identical to a complete *S.aureus* sequence from the sequence listing (e.g. when it comprises a sequence listing with <100% sequence identity thereto, or when it comprises a fragment thereof) it is preferred in each individual instance that the antigen can elicit an antibody which recognises the respective complete *S.aureus* sequence.

Mutant bacteria

5 [0459] The invention also provides a *S.aureus* bacterium in which one or more of the antigens from the various antigen groups of the invention has/have been knocked out. Techniques for producing knockout bacteria are well known, and knockout *S.aureus* strains have been reported. A knockout mutation may be situated in the coding region of the gene or may lie within its transcriptional control regions (e.g. within its promoter). A knockout mutation will reduce the level of mRNA encoding the antigen to <1% of that produced by the wild-type bacterium, preferably <0.5%, more preferably <0.1 %, and most preferably to 0%.

10 [0460] The invention also provides a *S.aureus* in which one or more of the antigens from the various antigen groups of the invention has a mutation which inhibits its activity. The gene encoding the antigen will have a mutation that changes the encoded amino acid sequence. Mutation may involve deletion, substitution, and/or insertion, any of which may be involve one or more amino acids.

[0461] The invention also provides a bacterium, such as a *S.aureus* bacterium, which hyper-expresses an antigen of the invention.

15 [0462] The invention also provides a bacterium, such as a *S.aureus* bacterium, that constitutively expresses an antigen of the invention. The invention also provides a meningococcus comprising a gene encoding an antigen of the invention, wherein the gene is under the control of an inducible promoter.

Immunogenic compositions and medicaments

20 [0463] Immunogenic compositions of the invention may be useful as vaccines. Vaccines according to the invention may either be prophylactic (i.e. to prevent infection) or therapeutic (i.e. to treat infection), but will typically be prophylactic.

[0464] Compositions may thus be pharmaceutically acceptable. They will usually include components in addition to the antigens e.g. they typically include one or more pharmaceutical carrier(s) and/or excipient(s). A thorough discussion of such components is available in reference 273.

25 [0465] Compositions will generally be administered to a mammal in aqueous form. Prior to administration, however, the composition may have been in a non-aqueous form. For instance, although some vaccines are manufactured in aqueous form, then filled and distributed and administered also in aqueous form, other vaccines are lyophilised during manufacture and are reconstituted into an aqueous form at the time of use. Thus a composition of the invention may be dried, such as a lyophilised formulation.

30 [0466] The composition may include preservatives such as thiomersal or 2-phenoxyethanol. It is preferred, however, that the vaccine should be substantially free from (i. e. less than 5 µg/ml) mercurial material e.g. thiomersal-free. Vaccines containing no mercury are more preferred. Preservative-free vaccines are particularly preferred.

[0467] To improve thermal stability, a composition may include a temperature protective agent. Further details of such agents are provided below.

35 [0468] To control tonicity, it is preferred to include a physiological salt, such as a sodium salt. Sodium chloride (NaCl) is preferred, which may be present at between 1 and 20 mg/ml e.g. about 10±2mg/ml NaCl. Other salts that may be present include potassium chloride, potassium dihydrogen phosphate, disodium phosphate dehydrate, magnesium chloride, calcium chloride, etc.

40 [0469] Compositions will generally have an osmolality of between 200 mOsm/kg and 400 mOsm/kg, preferably between 240-360 mOsm/kg, and will more preferably fall within the range of 290-310 mOsm/kg.

[0470] Compositions may include one or more buffers. Typical buffers include: a phosphate buffer; a Tris buffer; a borate buffer; a succinate buffer; a histidine buffer (particularly with an aluminum hydroxide adjuvant); or a citrate buffer. Buffers will typically be included in the 5-20mM range.

45 [0471] The pH of a composition will generally be between 5.0 and 8.1, and more typically between 6.0 and 8.0 e.g. 6.5 and 7.5, or between 7.0 and 7.8.

[0472] The composition is preferably sterile. The composition is preferably non-pyrogenic e.g. containing <1 EU (endotoxin unit, a standard measure) per dose, and preferably <0.1 EU per dose. The composition is preferably gluten free.

50 [0473] The composition may include material for a single immunisation, or may include material for multiple immunisations (i.e. a 'multidose' kit). The inclusion of a preservative is preferred in multidose arrangements. As an alternative (or in addition) to including a preservative in multidose compositions, the compositions may be contained in a container having an aseptic adaptor for removal of material.

[0474] Human vaccines are typically administered in a dosage volume of about 0.5ml, although a half dose (i.e. about 0.25ml) may be administered to children.

55 [0475] Immunogenic compositions of the invention may also comprise one or more immunoregulatory agents. Preferably, one or more of the immunoregulatory agents include one or more adjuvants. The adjuvants may include a TH1 adjuvant and/or a TH2 adjuvant, further discussed below.

[0476] Thus the invention provides an immunogenic composition comprising a combination of:

- (1) one or more antigen(s) selected from the first, second, third and fourth antigen groups (as defined above); and
 (2) an adjuvant, such as an aluminium hydroxide adjuvant (for example, one or more antigens may be adsorbed to aluminium hydroxide).

5 **[0477]** For instance, the invention provides an immunogenic composition comprising a combination of a sta006 antigen and an adjuvant, such as an aluminium hydroxide adjuvant. Similarly, the invention provides an immunogenic composition comprising a combination of a sta011 antigen and an adjuvant, such as an aluminium hydroxide adjuvant. These compositions are ideally buffered e.g. with a histidine buffer.

[0478] Adjuvants which may be used in compositions of the invention include, but are not limited to:

10 *A. Mineral-containing compositions*

[0479] Mineral containing compositions suitable for use as adjuvants in the invention include mineral salts, such as aluminium salts and calcium salts (or mixtures thereof). Calcium salts include calcium phosphate (e.g. the "CAP" particles disclosed in ref. 80). Aluminum salts include hydroxides, phosphates, sulfates, etc., with the salts taking any suitable form (e.g. gel, crystalline, amorphous, etc.). Adsorption to these salts is preferred (e.g. all antigens may be adsorbed). The mineral containing compositions may also be formulated as a particle of metal salt [81].

[0480] The adjuvants known as aluminium hydroxide and aluminum phosphate may be used. These names are conventional, but are used for convenience only, as neither is a precise description of the actual chemical compound which is present (e.g. see chapter 9 of reference 82)). The invention can use any of the "hydroxide" or "phosphate" adjuvants that are in general use as adjuvants. The adjuvants known as "aluminium hydroxide" are typically aluminium oxyhydroxide salts, which are usually at least partially crystalline. The adjuvants known as "aluminium phosphate" are typically aluminium hydroxyphosphates, often also containing a small amount of sulfate (i. e. aluminium hydroxyphosphate sulfate). They may be obtained by precipitation, and the reaction conditions and concentrations during precipitation influence the degree of substitution of phosphate for hydroxyl in the salt.

[0481] A fibrous morphology (e.g. as seen in transmission electron micrographs) is typical for aluminium hydroxide adjuvants. The pl of aluminium hydroxide adjuvants is typically about 11 i.e. the adjuvant itself has a positive surface charge at physiological pH. Adsorptive capacities of between 1.8-2.6 mg protein per mg Al⁺⁺⁺ at pH 7.4 have been reported for aluminium hydroxide adjuvants.

[0482] Aluminium phosphate adjuvants generally have a PO₄/Al molar ratio between 0.3 and 1.2, preferably between 0.8 and 1.2, and more preferably 0.95±0.1. The aluminium phosphate will generally be amorphous, particularly for hydroxyphosphate salts. A typical adjuvant is amorphous aluminium hydroxyphosphate with PO₄/Al molar ratio between 0.84 and 0.92, included at 0.6mg Al³⁺/ml. The aluminium phosphate will generally be particulate (e.g. plate-like morphology as seen in transmission electron micrographs). Typical diameters of the particles are in the range 0.5-20µm (e.g. about 5-10µm) after any antigen adsorption. Adsorptive capacities of between 0.7-1.5 mg protein per mg Al⁺⁺⁺ at pH 7.4 have been reported for aluminium phosphate adjuvants.

[0483] The point of zero charge (PZC) of aluminium phosphate is inversely related to the degree of substitution of phosphate for hydroxyl, and this degree of substitution can vary depending on reaction conditions and concentration of reactants used for preparing the salt by precipitation. PZC is also altered by changing the concentration of free phosphate ions in solution (more phosphate = more acidic PZC) or by adding a buffer such as a histidine buffer (makes PZC more basic). Aluminium phosphates used according to the invention will generally have a PZC of between 4.0 and 7.0, more preferably between 5.0 and 6.5 e.g. about 5.7.

[0484] As shown below, adsorption of *S.aureus* protein antigens (except IsdA, Sta019 and Sta073) to an aluminium hydroxide adjuvant is advantageous, particularly in a multi-protein combination (in which all antigens may be adsorbed). A histidine buffer can usefully be included in such adjuvanted compositions.

[0485] Suspensions of aluminium salts used to prepare compositions of the invention may contain a buffer (e.g. a phosphate or a histidine or a Tris buffer), but this is not always necessary. The suspensions are preferably sterile and pyrogen-free. A suspension may include free aqueous phosphate ions e.g. present at a concentration between 1.0 and 20 mM, preferably between 5 and 15 mM, and more preferably about 10 mM. The suspensions may also comprise sodium chloride.

[0486] The invention can use a mixture of both an aluminium hydroxide and an aluminium phosphate. In this case there may be more aluminium phosphate than hydroxide e.g. a weight ratio of at least 2:1 e.g. ≥5:1, ≥6:1, ≥7:1, ≥8:1, ≥9:1, etc.

[0487] The concentration of Al⁺⁺⁺ in a composition for administration to a patient is preferably less than 10mg/ml e.g. ≤5 mg/ml, ≤4 mg/ml, ≤3 mg/ml, ≤2 mg/ml, ≤1 mg/ml, etc. A preferred range is between 0.3 and 1mg/ml. A maximum of 0.85mg/dose is preferred.

B. Oil Emulsions

[0488] Oil emulsion compositions suitable for use as adjuvants in the invention include squalene-water emulsions, such as MF59 [Chapter 10 of ref. 82; see also ref. 83] (5% Squalene, 0.5% Tween 80, and 0.5% Span 85, formulated into submicron particles using a microfluidizer). Complete Freund's adjuvant (CFA) and incomplete Freund's adjuvant (IFA) may also be used.

[0489] Various oil-in-water emulsion adjuvants are known, and they typically include at least one oil and at least one surfactant, with the oil(s) and surfactant(s) being biodegradable (metabolizable) and biocompatible. The oil droplets in the emulsion are generally less than 5 μm in diameter, and ideally have a sub-micron diameter, with these small sizes being achieved with a microfluidiser to provide stable emulsions. Droplets with a size less than 220nm are preferred as they can be subjected to filter sterilization.

[0490] The emulsion can comprise oils such as those from an animal (such as fish) or vegetable source. Sources for vegetable oils include nuts, seeds and grains. Peanut oil, soybean oil, coconut oil, and olive oil, the most commonly available, exemplify the nut oils. Jojoba oil can be used e.g. obtained from the jojoba bean. Seed oils include safflower oil, cottonseed oil, sunflower seed oil, sesame seed oil and the like. In the grain group, corn oil is the most readily available, but the oil of other cereal grains such as wheat, oats, rye, rice, teff, triticale and the like may also be used. 6-10 carbon fatty acid esters of glycerol and 1,2-propanediol, while not occurring naturally in seed oils, may be prepared by hydrolysis, separation and esterification of the appropriate materials starting from the nut and seed oils. Fats and oils from mammalian milk are metabolizable and may therefore be used in the practice of this invention. The procedures for separation, purification, saponification and other means necessary for obtaining pure oils from animal sources are well known in the art. Most fish contain metabolizable oils which may be readily recovered. For example, cod liver oil, shark liver oils, and whale oil such as spermaceti exemplify several of the fish oils which may be used herein. A number of branched chain oils are synthesized biochemically in 5-carbon isoprene units and are generally referred to as terpenoids. Shark liver oil contains a branched, unsaturated terpenoids known as squalene, 2,6,10,15,19,23-hexamethyl-2,6,10,14,18,22-tetracosahexaene, which is particularly preferred herein. Squalane, the saturated analog to squalene, is also a preferred oil. Fish oils, including squalene and squalane, are readily available from commercial sources or may be obtained by methods known in the art. Other preferred oils are the tocopherols (see below). Mixtures of oils can be used.

[0491] Surfactants can be classified by their 'HLB' (hydrophilic/lipophile balance). Preferred surfactants of the invention have a HLB of at least 10, preferably at least 15, and more preferably at least 16. The invention can be used with surfactants including, but not limited to: the polyoxyethylene sorbitan esters surfactants (commonly referred to as the Tweens), especially polysorbate 20 and polysorbate 80; copolymers of ethylene oxide (EO), propylene oxide (PO), and/or butylene oxide (BO), sold under the DOWFAX™ tradename, such as linear EO/PO block copolymers; octoxynols, which can vary in the number of repeating ethoxy (oxy-1,2-ethanediyl) groups, with octoxynol-9 (Triton X-100, or t-octylphenoxypolyethoxyethanol) being of particular interest; (octylphenoxy)polyethoxyethanol (IGEPAL CA-630/NP-40); phospholipids such as phosphatidylcholine (lecithin); nonylphenol ethoxylates, such as the Tergitol™ NP series; polyoxyethylene fatty ethers derived from lauryl, cetyl, stearyl and oleyl alcohols (known as Brij surfactants), such as triethyleneglycol monolauryl ether (Brij 30); and sorbitan esters (commonly known as the SPANs), such as sorbitan trioleate (Span 85) and sorbitan monolaurate. Non-ionic surfactants are preferred. Preferred surfactants for including in the emulsion are Tween 80 (polyoxyethylene sorbitan monooleate), Span 85 (sorbitan trioleate), lecithin and Triton X-100.

[0492] Mixtures of surfactants can be used e.g. Tween 80/Span 85 mixtures. A combination of a polyoxyethylene sorbitan ester such as polyoxyethylene sorbitan monooleate (Tween 80) and an octoxynol such as t-octylphenoxypolyethoxyethanol (Triton X-100) is also suitable. Another useful combination comprises laureth 9 plus a polyoxyethylene sorbitan ester and/or an octoxynol.

[0493] Preferred amounts of surfactants (% by weight) are: polyoxyethylene sorbitan esters (such as Tween 80) 0.01 to 1%, in particular about 0.1%; octyl- or nonylphenoxy polyoxyethanols (such as Triton X-100, or other detergents in the Triton series) 0.001 to 0.1%, in particular 0.005 to 0.02%; polyoxyethylene ethers (such as laureth 9) 0.1 to 20%, preferably 0.1 to 10% and in particular 0.1 to 1% or about 0.5%.

[0494] Preferred emulsion adjuvants have an average droplets size of $<1 \mu\text{m}$ e.g. $\leq 750\text{nm}$, $\leq 500\text{nm}$, $\leq 400\text{nm}$, $\leq 300\text{nm}$, $\leq 250\text{nm}$, $\leq 220\text{nm}$, $\leq 200\text{nm}$, or smaller. These droplet sizes can conveniently be achieved by techniques such as microfluidisation.

[0495] Specific oil-in-water emulsion adjuvants useful with the invention include, but are not limited to:

- A submicron emulsion of squalene, Tween 80, and Span 85. The composition of the emulsion by volume can be about 5% squalene, about 0.5% polysorbate 80 and about 0.5% Span 85. In weight terms, these ratios become 4.3% squalene, 0.5% polysorbate 80 and 0.48% Span 85. This adjuvant is known as 'MF59' [84-86], as described in more detail in Chapter 10 of ref. 87 and chapter 12 of ref. 88. The MF59 emulsion advantageously includes citrate ions e.g. 10mM sodium citrate buffer.

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● An emulsion of squalene, a tocopherol, and polysorbate 80 (Tween 80). The emulsion may include phosphate buffered saline. It may also include Span 85 (*e.g.* at 1%) and/or lecithin. These emulsions may have from 2 to 10% squalene, from 2 to 10% tocopherol and from 0.3 to 3% Tween 80, and the weight ratio of squalene:tocopherol is preferably ≤ 1 as this provides a more stable emulsion. Squalene and Tween 80 may be present volume ratio of about 5:2 or at a weight ratio of about 11:5. One such emulsion can be made by dissolving Tween 80 in PBS to give a 2% solution, then mixing 90ml of this solution with a mixture of (5g of DL- α -tocopherol and 5ml squalene), then microfluidising the mixture. The resulting emulsion may have submicron oil droplets *e.g.* with an average diameter of between 100 and 250nm, preferably about 180nm. The emulsion may also include a 3-de-O-acylated monophosphoryl lipid A (3d-MPL). Another useful emulsion of this type may comprise, per human dose, 0.5-10 mg squalene, 0.5-11 mg tocopherol, and 0.1-4 mg polysorbate 80 [89].

● An emulsion of squalene, a tocopherol, and a Triton detergent (*e.g.* Triton X-100). The emulsion may also include a 3d-MPL (see below). The emulsion may contain a phosphate buffer.

● An emulsion comprising a polysorbate (*e.g.* polysorbate 80), a Triton detergent (*e.g.* Triton X-100) and a tocopherol (*e.g.* an α -tocopherol succinate). The emulsion may include these three components at a mass ratio of about 75:11:10 (*e.g.* 750 μ g/ml polysorbate 80, 110 μ g/ml Triton X-100 and 100 μ g/ml α -tocopherol succinate), and these concentrations should include any contribution of these components from antigens. The emulsion may also include squalene. The emulsion may also include a 3d-MPL (see below). The aqueous phase may contain a phosphate buffer.

● An emulsion of squalane, polysorbate 80 and poloxamer 401 ("Pluronic™ L121"). The emulsion can be formulated in phosphate buffered saline, pH 7.4. This emulsion is a useful delivery vehicle for muramyl dipeptides, and has been used with threonyl-MDP in the "SAF-1" adjuvant [90] (0.05-1% Thr-MDP, 5% squalane, 2.5% Pluronic L121 and 0.2% polysorbate 80). It can also be used without the Thr-MDP, as in the "AF" adjuvant [91] (5% squalane, 1.25% Pluronic L121 and 0.2% polysorbate 80). Microfluidisation is preferred.

● An emulsion comprising squalene, an aqueous solvent, a polyoxyethylene alkyl ether hydrophilic nonionic surfactant (*e.g.* polyoxyethylene (12) cetostearyl ether) and a hydrophobic nonionic surfactant (*e.g.* a sorbitan ester or mannide ester, such as sorbitan monoleate or 'Span 80'). The emulsion is preferably thermoreversible and/or has at least 90% of the oil droplets (by volume) with a size less than 200 nm [92]. The emulsion may also include one or more of: alditol; a cryoprotective agent (*e.g.* a sugar, such as dodecylmaltoside and/or sucrose); and/or an alkylpolyglycoside. The emulsion may include a TLR4 agonist [93]. Such emulsions may be lyophilized.

● An emulsion of squalene, poloxamer 105 and Abil-Care [94]. The final concentration (weight) of these components in adjuvanted vaccines are 5% squalene, 4% poloxamer 105 (pluronic polyol) and 2% Abil-Care 85 (Bis-PEG/PPG-16/16 PEG/PPG-16/16 dimethicone; caprylic/capric triglyceride).

● An emulsion having from 0.5-50% of an oil, 0.1-10% of a phospholipid, and 0.05-5% of a non-ionic surfactant. As described in reference 95, preferred phospholipid components are phosphatidylcholine, phosphatidylethanolamine, phosphatidylserine, phosphatidylinositol, phosphatidylglycerol, phosphatidic acid, sphingomyelin and cardiolipin. Submicron droplet sizes are advantageous.

● A submicron oil-in-water emulsion of a non-metabolisable oil (such as light mineral oil) and at least one surfactant (such as lecithin, Tween 80 or Span 80). Additives may be included, such as QuilA saponin, cholesterol, a saponin-lipophile conjugate (such as GPI-0100, described in reference 96, produced by addition of aliphatic amine to desacylsaponin via the carboxyl group of glucuronic acid), dimethyldioctadecylammonium bromide and/or N,N-dioctadecyl-N,N-bis (2-hydroxyethyl)propanediamine.

● An emulsion in which a saponin (*e.g.* QuilA or QS21) and a sterol (*e.g.* a cholesterol) are associated as helical micelles [97].

● An emulsion comprising a mineral oil, a non-ionic lipophilic ethoxylated fatty alcohol, and a non-ionic hydrophilic surfactant (*e.g.* an ethoxylated fatty alcohol and/or polyoxyethylene-polyoxypropylene block copolymer) [98].

● An emulsion comprising a mineral oil, a non-ionic hydrophilic ethoxylated fatty alcohol, and a non-ionic lipophilic surfactant (*e.g.* an ethoxylated fatty alcohol and/or polyoxyethylene-polyoxypropylene block copolymer) [98].

[0496] In some embodiments an emulsion may be mixed with antigen extemporaneously, at the time of delivery, and

thus the adjuvant and antigen may be kept separately in a packaged or distributed vaccine, ready for final formulation at the time of use. In other embodiments an emulsion is mixed with antigen during manufacture, and thus the composition is packaged in a liquid adjuvanted form,. The antigen will generally be in an aqueous form, such that the vaccine is finally prepared by mixing two liquids. The volume ratio of the two liquids for mixing can vary (e.g. between 5:1 and 1:5) but is generally about 1:1. Where concentrations of components are given in the above descriptions of specific emulsions, these concentrations are typically for an undiluted composition, and the concentration after mixing with an antigen solution will thus decrease.

[0497] Where a composition includes a tocopherol, any of the α , β , γ , δ , ϵ or ξ tocopherols can be used, but α -tocopherols are preferred. The tocopherol can take several forms e.g. different salts and/or isomers. Salts include organic salts, such as succinate, acetate, nicotinate, etc. D- α -tocopherol and DL- α -tocopherol can both be used. Tocopherols are advantageously included in vaccines for use in elderly patients (e.g. aged 60 years or older) because vitamin E has been reported to have a positive effect on the immune response in this patient group [99]. They also have antioxidant properties that may help to stabilize the emulsions [100]. A preferred α -tocopherol is DL- α -tocopherol, and the preferred salt of this tocopherol is the succinate. The succinate salt has been found to cooperate with TNF-related ligands *in vivo*.

C. Saponin formulations [chapter 22 of ref. 82]

[0498] Saponin formulations may also be used as adjuvants in the invention. Saponins are a heterogeneous group of sterol glycosides and triterpenoid glycosides that are found in the bark, leaves, stems, roots and even flowers of a wide range of plant species. Saponin from the bark of the *Quillaia saponaria* Molina tree have been widely studied as adjuvants. Saponin can also be commercially obtained from *Smilax ornata* (sarsapilla), *Gypsophilla paniculata* (brides veil), and *Saponaria officianalis* (soap root). Saponin adjuvant formulations include purified formulations, such as QS21, as well as lipid formulations, such as ISCOMs. QS21 is marketed as Stimulon™.

[0499] Saponin compositions have been purified using HPLC and RP-HPLC. Specific purified fractions using these techniques have been identified, including QS7, QS17, QS18, QS21, QH-A, QH-B and QH-C. Preferably, the saponin is QS21. A method of production of QS21 is disclosed in ref. 101. Saponin formulations may also comprise a sterol, such as cholesterol [102].

[0500] Combinations of saponins and cholesterol can be used to form unique particles called immunostimulating complexes (ISCOMs) [chapter 23 of ref. 82]. ISCOMs typically also include a phospholipid such as phosphatidylethanolamine or phosphatidylcholine. Any known saponin can be used in ISCOMs. Preferably, the ISCOM includes one or more of QuilA, QHA & QHC. ISCOMs are further described in refs. 102-104. Optionally, the ISCOMS may be devoid of additional detergent [105].

[0501] A review of the development of saponin based adjuvants can be found in refs. 106 & 107.

D. Virosomes and virus-like particles

[0502] Virosomes and virus-like particles (VLPs) can also be used as adjuvants in the invention. These structures generally contain one or more proteins from a virus optionally combined or formulated with a phospholipid. They are generally non-pathogenic, non-replicating and generally do not contain any of the native viral genome. The viral proteins may be recombinantly produced or isolated from whole viruses. These viral proteins suitable for use in virosomes or VLPs include proteins derived from influenza virus (such as HA or NA), Hepatitis B virus (such as core or capsid proteins), Hepatitis E virus, measles virus, Sindbis virus, Rotavirus, Foot-and-Mouth Disease virus, Retrovirus, Norwalk virus, human Papilloma virus, HIV, RNA-phages, QB-phage (such as coat proteins), GA-phage, fr-phage, AP205 phage, and Ty (such as retrotransposon Ty protein pl). VLPs are discussed further in refs. 108-113. Virosomes are discussed further in, for example, ref. 114

E. Bacterial or microbial derivatives

[0503] Adjuvants suitable for use in the invention include bacterial or microbial derivatives such as non-toxic derivatives of enterobacterial lipopolysaccharide (LPS), Lipid A derivatives, immunostimulatory oligonucleotides and ADP-ribosylating toxins and detoxified derivatives thereof.

[0504] Non-toxic derivatives of LPS include monophosphoryl lipid A (MPL) and 3-O-deacylated MPL (3dMPL). 3dMPL is a mixture of 3 de-O-acylated monophosphoryl lipid A with 4, 5 or 6 acylated chains. A preferred "small particle" form of 3 De-O-acylated monophosphoryl lipid A is disclosed in ref. 115. Such "small particles" of 3dMPL are small enough to be sterile filtered through a 0.22 μ m membrane [115]. Other non-toxic LPS derivatives include monophosphoryl lipid A mimics, such as aminoalkyl glucosaminide phosphate derivatives e.g. RC-529 [116,117].

[0505] Lipid A derivatives include derivatives of lipid A from *Escherichia coli* such as OM-174. OM-174 is described for example in refs. 118 & 119.

[0506] Immunostimulatory oligonucleotides suitable for use as adjuvants in the invention include nucleotide sequences containing a CpG motif (a dinucleotide sequence containing an unmethylated cytosine linked by a phosphate bond to a guanosine). Double-stranded RNAs and oligonucleotides containing palindromic or poly(dG) sequences have also been shown to be immunostimulatory.

[0507] The CpG's can include nucleotide modifications/analogs such as phosphorothioate modifications and can be double-stranded or single-stranded. References 120, 121 and 122 disclose possible analog substitutions *e.g.* replacement of guanosine with 2'-deoxy-7-deazaguanosine. The adjuvant effect of CpG oligonucleotides is further discussed in refs. 123-128.

[0508] The CpG sequence may be directed to TLR9, such as the motif GTCGTT or TTCGTT [129]. The CpG sequence may be specific for inducing a Th1 immune response, such as a CpG-A ODN, or it may be more specific for inducing a B cell response, such as a CpG-B ODN. CpG-A and CpG-B ODNs are discussed in refs. 130-132. Preferably, the CpG is a CpG-A ODN.

[0509] Preferably, the CpG oligonucleotide is constructed so that the 5' end is accessible for receptor recognition. Optionally, two CpG oligonucleotide sequences may be attached at their 3' ends to form "immunomers". See, for example, refs. 129 & 133-135.

[0510] A useful CpG adjuvant is CpG7909, also known as ProMune™ (Coley Pharmaceutical Group, Inc.). Another is CpG1826. As an alternative, or in addition, to using CpG sequences, TpG sequences can be used [136], and these oligonucleotides may be free from unmethylated CpG motifs. The immunostimulatory oligonucleotide may be pyrimidine-rich. For example, it may comprise more than one consecutive thymidine nucleotide (*e.g.* TTTT, as disclosed in ref. 136), and/or it may have a nucleotide composition with >25% thymidine (*e.g.* >35%, >40%, >50%, >60%, >80%, *etc.*). For example, it may comprise more than one consecutive cytosine nucleotide (*e.g.* CCCC, as disclosed in ref. 136), and/or it may have a nucleotide composition with >25% cytosine (*e.g.* >35%, >40%, >50%, >60%, >80%, *etc.*). These oligonucleotides may be free from unmethylated CpG motifs. Immunostimulatory oligonucleotides will typically comprise at least 20 nucleotides. They may comprise fewer than 100 nucleotides.

[0511] A particularly useful adjuvant based around immunostimulatory oligonucleotides is known as IC-31™ [137]. Thus an adjuvant used with the invention may comprise a mixture of (i) an oligonucleotide (*e.g.* between 15-40 nucleotides) including at least one (and preferably multiple) CpI motifs (*i.e.* a cytosine linked to an inosine to form a dinucleotide), and (ii) a polycationic polymer, such as an oligopeptide (*e.g.* between 5-20 amino acids) including at least one (and preferably multiple) Lys-Arg-Lys tripeptide sequence(s). The oligonucleotide may be a deoxynucleotide comprising 26-mer sequence 5'-(IC)₁₃-3' (SEQ ID NO: 175). The polycationic polymer may be a peptide comprising 11-mer amino acid sequence KKLKLLKLLK (SEQ ID NO: 176). The oligonucleotide and polymer can form complexes *e.g.* as disclosed in references 138 & 139.

[0512] Bacterial ADP-ribosylating toxins and detoxified derivatives thereof may be used as adjuvants in the invention. Preferably, the protein is derived from *E. coli* (*E. coli* heat labile enterotoxin "LT"), cholera ("CT"), or pertussis ("PT"). The use of detoxified ADP-ribosylating toxins as mucosal adjuvants is described in ref. 140 and as parenteral adjuvants in ref. 141. The toxin or toxoid is preferably in the form of a holotoxin, comprising both A and B subunits. Preferably, the A subunit contains a detoxifying mutation; preferably the B subunit is not mutated. Preferably, the adjuvant is a detoxified LT mutant such as LT-K63, LT-R72, and LT-G192. The use of ADP-ribosylating toxins and detoxified derivatives thereof, particularly LT-K63 and LT-R72, as adjuvants can be found in refs. 142-149. A useful CT mutant is or CT-E29H [150]. Numerical reference for amino acid substitutions is preferably based on the alignments of the A and B subunits of ADP-ribosylating toxins set forth in ref. 151, specifically incorporated herein by reference in its entirety.

F. Human immunomodulators

[0513] Human immunomodulators suitable for use as adjuvants in the invention include cytokines, such as interleukins (*e.g.* IL-1, IL-2, IL-4, IL-5, IL-6, IL-7, IL-12 [152], *etc.*) [153], interferons (*e.g.* interferon- γ), macrophage colony stimulating factor, and tumor necrosis factor. A preferred immunomodulator is IL-12.

G. Bioadhesives and Mucoadhesives

[0514] Bioadhesives and mucoadhesives may also be used as adjuvants in the invention. Suitable bioadhesives include esterified hyaluronic acid microspheres [154] or mucoadhesives such as cross-linked derivatives of poly(acrylic acid), polyvinyl alcohol, polyvinyl pyrrolidone, polysaccharides and carboxymethylcellulose. Chitosan and derivatives thereof may also be used as adjuvants in the invention [155].

H. Microparticles

[0515] Microparticles may also be used as adjuvants in the invention. Microparticles (*i.e.* a particle of ~100nm to ~

150µm in diameter, more preferably ~200nm to ~30µm in diameter, and most preferably ~500nm to ~10µm in diameter) formed from materials that are biodegradable and non-toxic (e.g. a poly(α-hydroxy acid), a polyhydroxybutyric acid, a polyorthoester, a polyanhydride, a polycaprolactone, etc.), with poly(lactide-co-glycolide) are preferred, optionally treated to have a negatively-charged surface (e.g. with SDS) or a positively-charged surface (e.g. with a cationic detergent, such as CTAB).

I. Liposomes (Chapters 13 & 14 of ref. 82)

[0516] Examples of liposome formulations suitable for use as adjuvants are described in refs. 156-158.

J. Polyoxyethylene ether and polyoxyethylene ester formulations

[0517] Adjuvants suitable for use in the invention include polyoxyethylene ethers and polyoxyethylene esters [159]. Such formulations further include polyoxyethylene sorbitan ester surfactants in combination with an octoxynol [160] as well as polyoxyethylene alkyl ethers or ester surfactants in combination with at least one additional non-ionic surfactant such as an octoxynol [161]. Preferred polyoxyethylene ethers are selected from the following group: polyoxyethylene-9-lauryl ether (laureth 9), polyoxyethylene-9-stearyl ether, polyoxyethylene-8-stearyl ether, polyoxyethylene-4-lauryl ether, polyoxyethylene-35-lauryl ether, and polyoxyethylene-23-lauryl ether.

K. Phosphazenes

[0518] A phosphazene, such as poly(di(carboxylatophenoxy)phosphazene) ("PCPP") as described, for example, in references 162 and 163, may be used.

L. Muramyl peptides

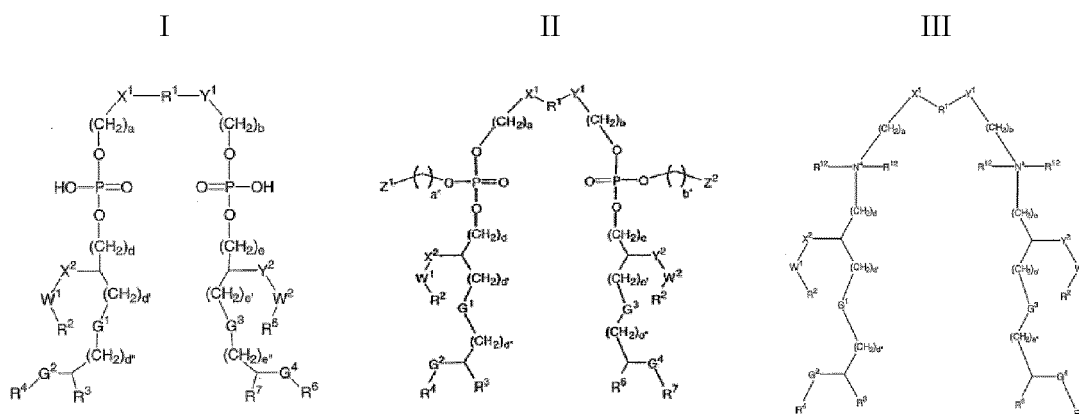
[0519] Examples of muramyl peptides suitable for use as adjuvants in the invention include N-acetyl-muramyl-L-threonyl-D-isoglutamine (thr-MDP), N-acetyl-normuramyl-L-alanyl-D-isoglutamine (nor-MDP), and N-acetylmuramyl-L-alanyl-D-isoglutaminyl-L-alanine-2-(1'-2'-dipalmitoyl-sn-glycero-3-hydroxyphosphoryloxy)-ethylamine MTP-PE.

M. Imidazoquinolone Compounds.

[0520] Examples of imidazoquinolone compounds suitable for use adjuvants in the invention include Imiquimod ("R-837") [164,165], Resiquimod ("R-848") [166], and their analogs; and salts thereof (e.g. the hydrochloride salts). Further details about immunostimulatory imidazoquinolines can be found in references 167 to 171.

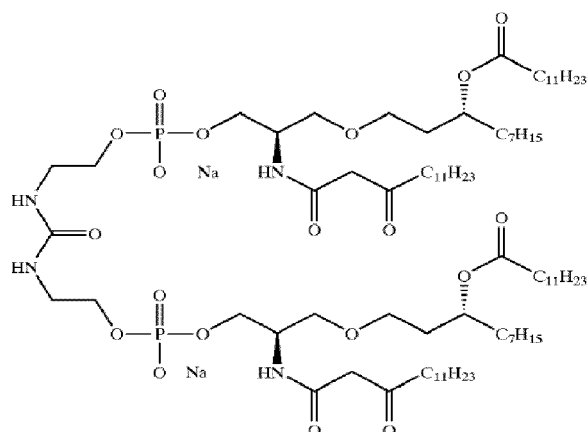
N. Substituted ureas

[0521] Substituted ureas useful as adjuvants include compounds of formula I, II or III, or salts thereof:



as defined in reference 172, such as 'ER 803058', 'ER 803732', 'ER 804053', 'ER 804058', 'ER 804059', 'ER 804442', 'ER 804680', 'ER 804764', 'ER 803022 or 'ER 804057' e.g.:

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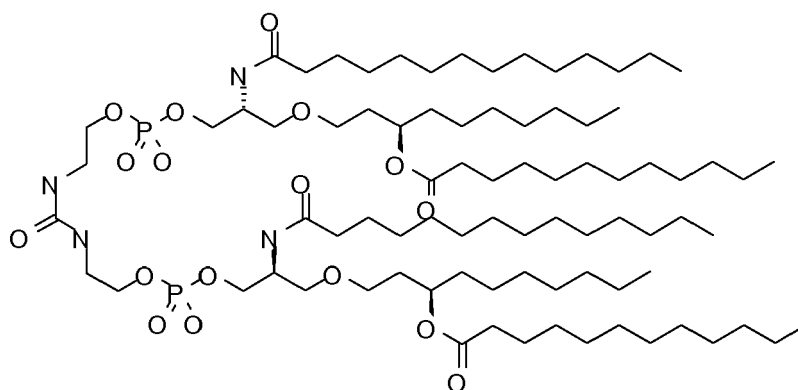


ER804057

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ER-803022:

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O. Further adjuvants

[0522] Further adjuvants that may be used with the invention include:

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- An aminoalkyl glucosaminide phosphate derivative, such as RC-529 [173,174].

[0523] Cyclic diguanylate ('c-di-GMP'), which has been reported as a useful adjuvant for *S.aureus* vaccines [175].

[0524] A thiosemicarbazone compound, such as those disclosed in reference 176. Methods of formulating, manufacturing, and screening for active compounds are also described in reference 176. The thiosemicarbazones are particularly effective in the stimulation of human peripheral blood mononuclear cells for the production of cytokines, such as TNF- α .

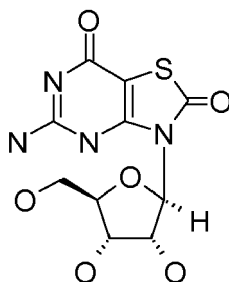
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[0525] A tryptanthrin compound, such as those disclosed in reference 177. Methods of formulating, manufacturing, and screening for active compounds are also described in reference 177. The thiosemicarbazones are particularly effective in the stimulation of human peripheral blood mononuclear cells for the production of cytokines, such as TNF- α .

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- A nucleoside analog, such as: (a) Isatorabine (ANA-245; 7-thia-8-oxoguanosine):

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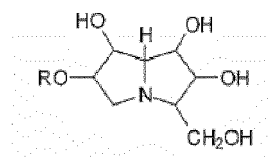


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and prodrugs thereof; (b) ANA975; (c) ANA-025-1; (d) ANA380; (e) the compounds disclosed in references 178 to 180 Loxoribine (7-allyl-8-oxoguanosine) [181].

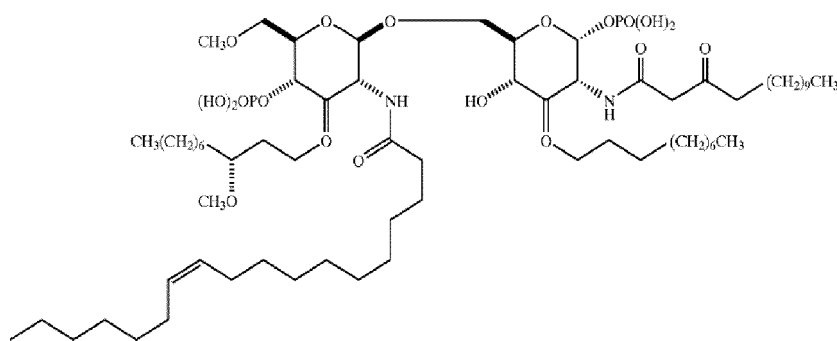
EP 2 510 947 A1

- Compounds disclosed in reference 182, including: Acylpiperazine compounds, Indoleione compounds, Tetrahydroisoquinoline (THIQ) compounds, Benzocyclodione compounds, Aminoazavinyl compounds, Aminobenzimidazole quinolinone (ABIQ) compounds [183,184], Hydrapthalamide compounds, Benzophenone compounds, Isoxazole compounds, Sterol compounds, Quinazolinone compounds, Pyrrole compounds [185], Anthraquinone compounds, Quinoxaline compounds, Triazine compounds, Pyrazalopyrimidine compounds, and Benzazole compounds [186].
- Compounds containing lipids linked to a phosphate-containing acyclic backbone, such as the TLR4 antagonist E5564 [187,188]:
- A polyoxidonium polymer [189,190] or other N-oxidized polyethylene-piperazine derivative.
- Methyl inosine 5'-monophosphate ("MIMP") [191].
- A polyhydroxylated pyrrolizidine compound [192], such as one having formula:



where R is selected from the group comprising hydrogen, straight or branched, unsubstituted or substituted, saturated or unsaturated acyl, alkyl (e.g. cycloalkyl), alkenyl, alkynyl and aryl groups, or a pharmaceutically acceptable salt or derivative thereof. Examples include, but are not limited to: casuarine, casuarine-6- α -D-glucopyranose, 3-*epi*-casuarine, 7-*epi*-casuarine, 3,7-*diepi*-casuarine, etc.

- A CDId ligand, such as an α -glycosylceramide [193-200] (e.g. α -galactosylceramide), phytosphingosine-containing α -glycosylceramides, OCH, KRN7000 [(2S,3S,4R)-1-O-(α -D-galactopyranosyl)-2-(N-hexacosanoylamino)-1,3,4-octadecanetriol], CRONY-101, 3"-O-sulfo-galactosylceramide, etc.
- A gamma inulin [201] or derivative thereof, such as algammulin.



Adjuvant combinations

[0526] The invention may also comprise combinations of one or more of the adjuvants identified above. For example, the following adjuvant compositions may be used in the invention: (1) a saponin and an oil-in-water emulsion [202]; (2) a saponin (e.g. QS21) + a non-toxic LPS derivative (e.g. 3dMPL) [203]; (3) a saponin (e.g. QS21) + a non-toxic LPS derivative (e.g. 3dMPL) + a cholesterol; (4) a saponin (e.g. QS21) + 3dMPL + IL-12 (optionally + a sterol) [204]; (5) combinations of 3dMPL with, for example, QS21 and/or oil-in-water emulsions [205]; (6) SAF, containing 10% squalene, 0.4% Tween 80™, 5% pluronic-block polymer L121, and thr-MDP, either microfluidized into a submicron emulsion or vortexed to generate a larger particle size emulsion. (7) Ribi™ adjuvant system (RAS), (Ribi Immunochem) containing 2% squalene, 0.2% Tween 80, and one or more bacterial cell wall components from the group consisting of monophosphorylipid A (MPL), trehalose dimycolate (TDM), and cell wall skeleton (CWS), preferably MPL + CWS (Detox™); and (8) one or more mineral salts (such as an aluminum salt) + a non-toxic derivative of LPS (such as 3dMPL).

[0527] Other substances that act as immunostimulating agents are disclosed in chapter 7 of ref. 82.

[0528] The use of an aluminium hydroxide and/or aluminium phosphate adjuvant is particularly preferred, and antigens are generally adsorbed to these salts. Calcium phosphate is another preferred adjuvant. Other preferred adjuvant combinations include combinations of Th1 and Th2 adjuvants such as CpG & alum or resiquimod & alum. A combination of aluminium phosphate and 3dMPL may be used.

[0529] The compositions of the invention may elicit both a cell mediated immune response as well as a humoral immune response. This immune response will preferably induce long lasting (e.g. neutralising) antibodies and a cell mediated immunity that can quickly respond upon exposure to pneumococcus.

[0530] Two types of T cells, CD4 and CD8 cells, are generally thought necessary to initiate and/or enhance cell mediated immunity and humoral immunity. CD8 T cells can express a CD8 co-receptor and are commonly referred to as Cytotoxic T lymphocytes (CTLs). CD8 T cells are able to recognize or interact with antigens displayed on MHC Class I molecules.

[0531] CD4 T cells can express a CD4 co-receptor and are commonly referred to as T helper cells. CD4 T cells are able to recognize antigenic peptides bound to MHC class II molecules. Upon interaction with a MHC class II molecule, the CD4 cells can secrete factors such as cytokines. These secreted cytokines can activate B cells, cytotoxic T cells, macrophages, and other cells that participate in an immune response. Helper T cells or CD4+ cells can be further divided into two functionally distinct subsets: TH1 phenotype and TH2 phenotypes which differ in their cytokine and effector function.

[0532] Activated TH1 cells enhance cellular immunity (including an increase in antigen-specific CTL production) and are therefore of particular value in responding to intracellular infections. Activated TH1 cells may secrete one or more of IL-2, IFN- γ , and TNF- β . A TH1 immune response may result in local inflammatory reactions by activating macrophages, NK (natural killer) cells, and CD8 cytotoxic T cells (CTLs). A TH1 immune response may also act to expand the immune response by stimulating growth of B and T cells with IL-12. TH1 stimulated B cells may secrete IgG2a.

[0533] Activated TH2 cells enhance antibody production and are therefore of value in responding to extracellular infections. Activated TH2 cells may secrete one or more of IL-4, IL-5, IL-6, and IL-10. A TH2 immune response may result in the production of IgG1, IgE, IgA and memory B cells for future protection.

[0534] An enhanced immune response may include one or more of an enhanced TH1 immune response and a TH2 immune response.

[0535] A TH1 immune response may include one or more of an increase in CTLs, an increase in one or more of the cytokines associated with a TH1 immune response (such as IL-2, IFN- γ , and TNF- β), an increase in activated macrophages, an increase in NK activity, or an increase in the production of IgG2a. Preferably, the enhanced TH1 immune response will include an increase in IgG2a production.

[0536] A TH1 immune response may be elicited using a TH1 adjuvant. A TH1 adjuvant will generally elicit increased levels of IgG2a production relative to immunization of the antigen without adjuvant. TH1 adjuvants suitable for use in the invention may include for example saponin formulations, virosomes and virus like particles, non-toxic derivatives of enterobacterial lipopolysaccharide (LPS), immunostimulatory oligonucleotides. Immunostimulatory oligonucleotides, such as oligonucleotides containing a CpG motif, are preferred TH1 adjuvants for use in the invention.

[0537] A TH2 immune response may include one or more of an increase in one or more of the cytokines associated with a TH2 immune response (such as IL-4, IL-5, IL-6 and IL-10), or an increase in the production of IgG1, IgE, IgA and memory B cells. Preferably, the enhanced TH2 immune response will include an increase in IgG1 production.

[0538] A TH2 immune response may be elicited using a TH2 adjuvant. A TH2 adjuvant will generally elicit increased levels of IgG1 production relative to immunization of the antigen without adjuvant. TH2 adjuvants suitable for use in the invention include, for example, mineral containing compositions, oil-emulsions, and ADP-ribosylating toxins and detoxified derivatives thereof. Mineral containing compositions, such as aluminium salts are preferred TH2 adjuvants for use in the invention.

[0539] Preferably, the invention includes a composition comprising a combination of a TH1 adjuvant and a TH2 adjuvant. Preferably, such a composition elicits an enhanced TH1 and an enhanced TH2 response, i.e., an increase in the production of both IgG1 and IgG2a production relative to immunization without an adjuvant. Still more preferably, the composition comprising a combination of a TH1 and a TH2 adjuvant elicits an increased TH1 and/or an increased TH2 immune response relative to immunization with a single adjuvant (i.e., relative to immunization with a TH1 adjuvant alone or immunization with a TH2 adjuvant alone).

[0540] The immune response may be one or both of a TH1 immune response and a TH2 response. Preferably, immune response provides for one or both of an enhanced TH1 response and an enhanced TH2 response.

[0541] The enhanced immune response may be one or both of a systemic and a mucosal immune response. Preferably, the immune response provides for one or both of an enhanced systemic and an enhanced mucosal immune response. Preferably the mucosal immune response is a TH2 immune response. Preferably, the mucosal immune response includes an increase in the production of IgA.

[0542] *S. aureus* infections can affect various areas of the body and so the compositions of the invention may be

prepared in various forms. For example, the compositions may be prepared as injectables, either as liquid solutions or suspensions. Solid forms suitable for solution in, or suspension in, liquid vehicles prior to injection can also be prepared (e.g. a lyophilised composition or a spray-freeze dried composition). The composition may be prepared for topical administration e.g. as an ointment, cream or powder. The composition may be prepared for oral administration e.g. as a tablet or capsule, as a spray, or as a syrup (optionally flavoured). The composition may be prepared for pulmonary administration e.g. as an inhaler, using a fine powder or a spray. The composition may be prepared as a suppository or pessary. The composition may be prepared for nasal, aural or ocular administration e.g. as drops. The composition may be in kit form, designed such that a combined composition is reconstituted just prior to administration to a patient. Such kits may comprise one or more antigens in liquid form and one or more lyophilised antigens.

[0543] Where a composition is to be prepared extemporaneously prior to use (e.g. where a component is presented in lyophilised form) and is presented as a kit, the kit may comprise two vials, or it may comprise one ready-filled syringe and one vial, with the contents of the syringe being used to reactivate the contents of the vial prior to injection.

[0544] Immunogenic compositions used as vaccines comprise an immunologically effective amount of antigen(s), as well as any other components, as needed. By 'immunologically effective amount', it is meant that the administration of that amount to an individual, either in a single dose or as part of a series, is effective for treatment or prevention. This amount varies depending upon the health and physical condition of the individual to be treated, age, the taxonomic group of individual to be treated (e.g. non-human primate, primate, etc.), the capacity of the individual's immune system to synthesise antibodies, the degree of protection desired, the formulation of the vaccine, the treating doctor's assessment of the medical situation, and other relevant factors. It is expected that the amount will fall in a relatively broad range that can be determined through routine trials. Where more than one antigen is included in a composition then two antigens may be present at the same dose as each other or at different doses.

[0545] As mentioned above, a composition may include a temperature protective agent, and this component may be particularly useful in adjuvanted compositions (particularly those containing a mineral adjuvant, such as an aluminium salt). As described in reference 206, a liquid temperature protective agent may be added to an aqueous vaccine composition to lower its freezing point e.g. to reduce the freezing point to below 0°C. Thus the composition can be stored below 0°C, but above its freezing point, to inhibit thermal breakdown. The temperature protective agent also permits freezing of the composition while protecting mineral salt adjuvants against agglomeration or sedimentation after freezing and thawing, and may also protect the composition at elevated temperatures e.g. above 40°C. A starting aqueous vaccine and the liquid temperature protective agent may be mixed such that the liquid temperature protective agent forms from 1-80% by volume of the final mixture. Suitable temperature protective agents should be safe for human administration, readily miscible/soluble in water, and should not damage other components (e.g. antigen and adjuvant) in the composition. Examples include glycerin, propylene glycol, and/or polyethylene glycol (PEG). Suitable PEGs may have an average molecular weight ranging from 200-20,000 Da. In a preferred embodiment, the polyethylene glycol can have an average molecular weight of about 300 Da ('PEG-300').

[0546] The invention provides an immunogenic composition comprising: (i) one or more antigen(s) selected from the first, second, third or fourth antigen groups; and (ii) a temperature protective agent. This composition may be formed by mixing (i) an aqueous composition comprising one or more antigen(s) selected from the first, second, third or fourth antigen groups, with (ii) a temperature protective agent. The mixture may then be stored e.g. below 0°C, from 0-20°C, from 20-35°C, from 35-55°C, or higher. It may be stored in liquid or frozen form. The mixture may be lyophilised. The composition may alternatively be formed by mixing (i) a dried composition comprising one or more antigen(s) selected from the first, second, third or fourth antigen groups, with (ii) a liquid composition comprising the temperature protective agent. Thus component (ii) can be used to reconstitute component (i).

Methods of treatment, and administration of the vaccine

[0547] The invention also provides a method for raising an immune response in a mammal comprising the step of administering an effective amount of a composition of the invention. The immune response is preferably protective and preferably involves antibodies and/or cell-mediated immunity. The method may raise a booster response.

[0548] The invention also provides at least two antigens of the invention for combined use as a medicament e.g. for use in raising an immune response in a mammal.

[0549] The invention also provides the use of at least two antigens of the invention in the manufacture of a medicament for raising an immune response in a mammal.

[0550] By raising an immune response in the mammal by these uses and methods, the mammal can be protected against *S.aureus* infection, including a nosocomial infection. More particularly, the mammal may be protected against a skin infection, pneumonia, meningitis, osteomyelitis endocarditis, toxic shock syndrome, and/or septicaemia.

[0551] The invention also provides a kit comprising a first component and a second component wherein neither the first component nor the second component is a composition of the invention as described above, but wherein the first component and the second component can be combined to provide a composition of the invention as described above.

The kit may further include a third component comprising one or more of the following: instructions, syringe or other delivery device, adjuvant, or pharmaceutically acceptable formulating solution.

[0552] The invention also provides a delivery device pre-filled with an immunogenic composition of the invention.

[0553] The mammal is preferably a human. Where the vaccine is for prophylactic use, the human is preferably a child (e.g. a toddler or infant) or a teenager; where the vaccine is for therapeutic use, the human is preferably a teenager or an adult. A vaccine intended for children may also be administered to adults e.g. to assess safety, dosage, immunogenicity, etc. Other mammals which can usefully be immunised according to the invention are cows, dogs, horses, and pigs.

[0554] One way of checking efficacy of therapeutic treatment involves monitoring *S.aureus* infection after administration of the compositions of the invention. One way of checking efficacy of prophylactic treatment involves monitoring immune responses, systemically (such as monitoring the level of IgG1 and IgG2a production) and/or mucosally (such as monitoring the level of IgA production), against the antigens in the compositions of the invention after administration of the composition. Typically, antigen-specific serum antibody responses are determined post-immunisation but pre-challenge whereas antigen-specific mucosal antibody responses are determined post-immunisation and post-challenge.

[0555] Another way of assessing the immunogenicity of the compositions of the present invention is to express the proteins recombinantly for screening patient sera or mucosal secretions by immunoblot and/or microarrays. A positive reaction between the protein and the patient sample indicates that the patient has mounted an immune response to the protein in question. This method may also be used to identify immunodominant antigens and/or epitopes within antigens.

[0556] The efficacy of vaccine compositions can also be determined *in vivo* by challenging animal models of *S.aureus* infection, e.g., guinea pigs or mice, with the vaccine compositions. In particular, there are three useful animal models for the study of *S.aureus* infectious disease, namely: (i) the murine abscess model [207], (ii) the murine lethal infection model [207] and (iii) the murine pneumonia model [208]. The abscess model looks at abscesses in mouse kidneys after intravenous challenge. The lethal infection model looks at the number of mice which survive after being infected by a normally-lethal dose of *S.aureus* by the intravenous or intraperitoneal route. The pneumonia model also looks at the survival rate, but uses intranasal infection. A useful vaccine may be effective in one or more of these models. For instance, for some clinical situations it may be desirable to protect against pneumonia, without needing to prevent hematic spread or to promote opsonisation; in other situations the main desire may be to prevent hematic spread. Different antigens, and different antigen combinations, may contribute to different aspects of an effective vaccine.

[0557] Compositions of the invention will generally be administered directly to a patient. Direct delivery may be accomplished by parenteral injection (e.g. subcutaneously, intraperitoneally, intravenously, intramuscularly, or to the interstitial space of a tissue), or mucosally, such as by rectal, oral (e.g. tablet, spray), vaginal, topical, transdermal or transcutaneous, intranasal, ocular, aural, pulmonary or other mucosal administration.

[0558] The invention may be used to elicit systemic and/or mucosal immunity, preferably to elicit an enhanced systemic and/or mucosal immunity.

[0559] Preferably the enhanced systemic and/or mucosal immunity is reflected in an enhanced TH1 and/or TH2 immune response. Preferably, the enhanced immune response includes an increase in the production of IgG1 and/or IgG2a and/or IgA.

[0560] Dosage can be by a single dose schedule or a multiple dose schedule. Multiple doses may be used in a primary immunisation schedule and/or in a booster immunisation schedule. In a multiple dose schedule the various doses may be given by the same or different routes e.g. a parenteral prime and mucosal boost, a mucosal prime and parenteral boost, etc. Multiple doses will typically be administered at least 1 week apart (e.g. about 2 weeks, about 3 weeks, about 4 weeks, about 6 weeks, about 8 weeks, about 10 weeks, about 12 weeks, about 16 weeks, etc.).

[0561] Vaccines prepared according to the invention may be used to treat both children and adults. Thus a human patient may be less than 1 year old, 1-5 years old, 5-15 years old, 15-55 years old, or at least 55 years old. Preferred patients for receiving the vaccines are the elderly (e.g. ≥ 50 years old, ≥ 60 years old, and preferably ≥ 65 years), the young (e.g. ≤ 5 years old), hospitalised patients, healthcare workers, armed service and military personnel, pregnant women, the chronically ill, or immunodeficient patients. The vaccines are not suitable solely for these groups, however, and may be used more generally in a population.

[0562] Vaccines produced by the invention may be administered to patients at substantially the same time as (e.g. during the same medical consultation or visit to a healthcare professional or vaccination centre) other vaccines e.g. at substantially the same time as an influenza vaccine, a measles vaccine, a mumps vaccine, a rubella vaccine, a MMR vaccine, a varicella vaccine, a MMRV vaccine, a diphtheria vaccine, a tetanus vaccine, a pertussis vaccine, a DTP vaccine, a conjugated *H.influenzae* type b vaccine, an inactivated poliovirus vaccine, a hepatitis B virus vaccine, a meningococcal conjugate vaccine (such as a tetravalent A-C-W135-Y vaccine), a respiratory syncytial virus vaccine, etc. Further non-staphylococcal vaccines suitable for co-administration may include one or more antigens listed on pages 33-46 of reference 51.

Nucleic acid immunisation

[0563] The immunogenic compositions described above include polypeptide antigens from *S.aureus*. In all cases, however, the polypeptide antigens can be replaced by nucleic acids (typically DNA) encoding those polypeptides, to give compositions, methods and uses based on nucleic acid immunisation. Nucleic acid immunisation is now a developed field (e.g. see references 209 to 216 etc.).

[0564] The nucleic acid encoding the immunogen is expressed *in vivo* after delivery to a patient and the expressed immunogen then stimulates the immune system. The active ingredient will typically take the form of a nucleic acid vector comprising: (i) a promoter; (ii) a sequence encoding the immunogen, operably linked to the promoter; and optionally (iii) a selectable marker. Preferred vectors may further comprise (iv) an origin of replication; and (v) a transcription terminator downstream of and operably linked to (ii). In general, (i) & (v) will be eukaryotic and (iii) & (iv) will be prokaryotic.

[0565] Preferred promoters are viral promoters e.g. from cytomegalovirus (CMV). The vector may also include transcriptional regulatory sequences (e.g. enhancers) in addition to the promoter and which interact functionally with the promoter. Preferred vectors include the immediate-early CMV enhancer/promoter, and more preferred vectors also include CMV intron A. The promoter is operably linked to a downstream sequence encoding an immunogen, such that expression of the immunogen-encoding sequence is under the promoter's control.

[0566] Where a marker is used, it preferably functions in a microbial host (e.g. in a prokaryote, in a bacteria, in a yeast). The marker is preferably a prokaryotic selectable marker (e.g. transcribed under the control of a prokaryotic promoter). For convenience, typical markers are antibiotic resistance genes.

[0567] The vector of the invention is preferably an autonomously replicating episomal or extrachromosomal vector, such as a plasmid.

[0568] The vector of the invention preferably comprises an origin of replication. It is preferred that the origin of replication is active in prokaryotes but not in eukaryotes.

[0569] Preferred vectors thus include a prokaryotic marker for selection of the vector, a prokaryotic origin of replication, but a eukaryotic promoter for driving transcription of the immunogen-encoding sequence. The vectors will therefore (a) be amplified and selected in prokaryotic hosts without polypeptide expression, but (b) be expressed in eukaryotic hosts without being amplified. This arrangement is ideal for nucleic acid immunization vectors.

[0570] The vector of the invention may comprise a eukaryotic transcriptional terminator sequence downstream of the coding sequence. This can enhance transcription levels. Where the coding sequence does not have its own, the vector of the invention preferably comprises a polyadenylation sequence. A preferred polyadenylation sequence is from bovine growth hormone.

[0571] The vector of the invention may comprise a multiple cloning site

[0572] In addition to sequences encoding the immunogen and a marker, the vector may comprise a second eukaryotic coding sequence. The vector may also comprise an IRES upstream of said second sequence in order to permit translation of a second eukaryotic polypeptide from the same transcript as the immunogen. Alternatively, the immunogen-coding sequence may be downstream of an IRES.

[0573] The vector of the invention may comprise unmethylated CpG motifs e.g. unmethylated DNA sequences which have in common a cytosine preceding a guanosine, flanked by two 5' purines and two 3' pyrimidines. In their unmethylated form these DNA motifs have been demonstrated to be potent stimulators of several types of immune cell.

[0574] Vectors may be delivered in a targeted way. Receptor-mediated DNA delivery techniques are described in, for example, references 217 to 222. Therapeutic compositions containing a nucleic acid are administered in a range of about 100ng to about 200mg of DNA for local administration in a gene therapy protocol. Concentration ranges of about 500 ng to about 50 mg, about 1µg to about 2 mg, about 5µg to about 500µg, and about 20µg to about 100µg of DNA can also be used during a gene therapy protocol. Factors such as method of action (e.g. for enhancing or inhibiting levels of the encoded gene product) and efficacy of transformation and expression are considerations which will affect the dosage required for ultimate efficacy. Where greater expression is desired over a larger area of tissue, larger amounts of vector or the same amounts re-administered in a successive protocol of administrations, or several administrations to different adjacent or close tissue portions may be required to effect a positive therapeutic outcome. In all cases, routine experimentation in clinical trials will determine specific ranges for optimal therapeutic effect.

[0575] Vectors can be delivered using gene delivery vehicles. The gene delivery vehicle can be of viral or non-viral origin (see generally references 223 to 226).

[0576] Viral-based vectors for delivery of a desired nucleic acid and expression in a desired cell are well known in the art. Exemplary viral-based vehicles include, but are not limited to, recombinant retroviruses (e.g. references 227 to 237), alphavirus-based vectors (e.g. Sindbis virus vectors, Semliki forest virus (ATCC VR-67; ATCC VR-1247), Ross River virus (ATCC VR-373; ATCC VR-1246) and Venezuelan equine encephalitis virus (ATCC VR-923; ATCC VR-1250; ATCC VR 1249; ATCC VR-532); hybrids or chimeras of these viruses may also be used), poxvirus vectors (e.g. vaccinia, fowlpox, canarypox, modified vaccinia Ankara, etc.), adenovirus vectors, and adeno-associated virus (AAV) vectors (e.g. see refs. 238 to 243). Administration of DNA linked to killed adenovirus [244] can also be employed.

[0577] Non-viral delivery vehicles and methods can also be employed, including, but not limited to, polycationic condensed DNA linked or unlinked to killed adenovirus alone [e.g. 244], ligand-linked DNA [245], eukaryotic cell delivery vehicles cells [e.g. refs. 246 to 250] and nucleic charge neutralization or fusion with cell membranes. Naked DNA can also be employed. Exemplary naked DNA introduction methods are described in refs. 251 and 252. Liposomes (e.g. immunoliposomes) that can act as gene delivery vehicles are described in refs. 253 to 257. Additional approaches are described in references 258 & 259.

[0578] Further non-viral delivery suitable for use includes mechanical delivery systems such as the approach described in ref. 259. Moreover, the coding sequence and the product of expression of such can be delivered through deposition of photopolymerized hydrogel materials or use of ionizing radiation [e.g. refs. 260 & 261]. Other conventional methods for gene delivery that can be used for delivery of the coding sequence include, for example, use of hand-held gene transfer particle gun [262] or use of ionizing radiation for activating transferred genes [260 & 261].

[0579] Delivery DNA using PLG {poly(lactide-co-glycolide)} microparticles is a particularly preferred method e.g. by adsorption to the microparticles, which are optionally treated to have a negatively-charged surface (e.g. treated with SDS) or a positively-charged surface (e.g. treated with a cationic detergent, such as CTAB).

S.epidermidis

[0580] Although the invention focuses on *S.aureus*, the inventors also realise that the sta006 and sta011 antigens have homologs in *S.epidermidis*. For example, SEQ ID NO: 234 is the 'iron (Fe+3) ABC superfamily ATP binding cassette transporter, binding protein' from *S.epidermidis* strain M23864:W1, with 73% identity to SEQ ID NO: 42 (sta006), and SEQ ID NO: 235 is the 'putative lipoprotein' from *S.epidermidis* strain RP62A, with 67% identity to SEQ ID NO: 47 (sta011). *S.epidermidis* is commonly present on human skin and can sometimes cause illness. Infection is usually associated with medical devices, such as catheters, and is a cause of nosocomial infections. The results disclosed herein for sta006 and sta011 against *S.aureus* suggest that the homologous proteins in *S. epidermidis* could be useful for immunising against this pathogen.

[0581] The invention provides an immunogenic composition comprising:

(i) a polypeptide comprising an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 234; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 234, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more); and/or

(ii) a polypeptide comprising an amino acid sequence: (a) having 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to SEQ ID NO: 235; and/or (b) comprising a fragment of at least 'n' consecutive amino acids of SEQ ID NO: 235, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250 or more).

[0582] The composition may also include an adjuvant. These compositions are particularly useful for immunising a mammal (including a human) against *S.epidermidis* infection.

[0583] Preferred fragments of (b) comprise an epitope from SEQ ID NO: 234 or 235, respectively. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of SEQ ID NO: 234/235 while retaining at least one epitope of SEQ ID NO: 234/235.

[0584] More generally, the invention provides the use of the sta006 and/or sta011 homolog from any *Staphylococcus* species for immunising a mammal against that species.

Antibodies

[0585] Antibodies against *S.aureus* antigens can be used for passive immunisation. Thus the invention provides an antibody which is specific for an antigen in the first, second, third or fourth antigen groups. The invention also provides the use of such antibodies in therapy. The invention also provides the use of such antibodies in the manufacture of a medicament. The invention also provides a method for treating a mammal comprising the step of administering an effective amount of an antibody of the invention. As described above for immunogenic compositions, these methods and uses allow a mammal to be protected against *S.aureus* infection.

[0586] The term "antibody" includes intact immunoglobulin molecules, as well as fragments thereof which are capable of binding an antigen. These include hybrid (chimeric) antibody molecules [263, 264]; F(ab')₂ and F(ab) fragments and Fv molecules; non-covalent heterodimers [265, 266]; single-chain Fv molecules (sFv) [267]; dimeric and trimeric antibody

fragment constructs; minibodies [268, 269]; humanized antibody molecules [270-272]; and any functional fragments obtained from such molecules, as well as antibodies obtained through non-conventional processes such as phage display. Preferably, the antibodies are monoclonal antibodies. Methods of obtaining monoclonal antibodies are well known in the art. Humanised or fully-human antibodies are preferred.

General

[0587] The practice of the present invention will employ, unless otherwise indicated, conventional methods of chemistry, biochemistry, molecular biology, immunology and pharmacology, within the skill of the art. Such techniques are explained fully in the literature. See, *e.g.*, references 273-280, *etc.*

[0588] "GI" numbering is used above. A GI number, or "GenInfo Identifier", is a series of digits assigned consecutively to each sequence record processed by NCBI when sequences are added to its databases. The GI number bears no resemblance to the accession number of the sequence record. When a sequence is updated (*e.g.* for correction, or to add more annotation or information) then it receives a new GI number. Thus the sequence associated with a given GI number is never changed.

[0589] Where the invention concerns an "epitope", this epitope may be a B-cell epitope and/or a T-cell epitope. Such epitopes can be identified empirically (*e.g.* using PEPSCAN [281,282] or similar methods), or they can be predicted (*e.g.* using the Jameson-Wolf antigenic index [283], matrix-based approaches [284], MAPITOPE [285], TEPITOPE [286,287], neural networks [288], OptiMer & EpiMer [289, 290], ADEPT [291], Tsites [292], hydrophilicity [293], antigenic index [294] or the methods disclosed in references 295-299, *etc.*). Epitopes are the parts of an antigen that are recognised by and bind to the antigen binding sites of antibodies or T-cell receptors, and they may also be referred to as "antigenic determinants".

[0590] Where an antigen "domain" is omitted, this may involve omission of a signal peptide, of a cytoplasmic domain, of a transmembrane domain, of an extracellular domain, *etc.*

[0591] The term "comprising" encompasses "including" as well as "consisting" *e.g.* a composition "comprising" X may consist exclusively of X or may include something additional *e.g.* X + Y.

[0592] The term "about" in relation to a numerical value x is optional and means, for example, $x \pm 10\%$.

[0593] References to a percentage sequence identity between two amino acid sequences means that, when aligned, that percentage of amino acids are the same in comparing the two sequences. This alignment and the percent homology or sequence identity can be determined using software programs known in the art, for example those described in section 7.7.18 of ref. 300. A preferred alignment is determined by the Smith-Waterman homology search algorithm using an affine gap search with a gap open penalty of 12 and a gap extension penalty of 2, BLOSUM matrix of 62. The Smith-Waterman homology search algorithm is disclosed in ref. 301.

BRIEF DESCRIPTION OF DRAWINGS

[0594]

Figure 1 shows bacterial counts (Log cfu/ml) after challenge of mice previously immunised with the indicated antigens.

Figures 2 to 4 show survival (%) after challenge of mice previously immunised with various mixtures of antigens over 14 days. In Figure 2, the six groups from SA-10-a are, from top to bottom at day 14. groups (i), (iii) & (iv) together, (ii), IsdB, then the negative control. In Figure 3, the six groups from SA-10-a are, from top to bottom at day 14. groups (i), (iii) & (iv) together, (ii), IsdB, then the negative control. In Figure 3, the six groups from SA-10-b are, from top to bottom at day 14.

groups (iii), (i), (iv), (ii) and IsdB together, then the negative control. In Figure 4, the six groups from SA-14 are, from top to bottom at day 14. groups (iv), (ii), (i), (iii), negative control, and IsdB.

Figure 5 shows collected data on mouse survival from four different experiments after challenge of mice previously immunised with various compositions (PBS negative control; IsdB antigen; and "Combo-1" and "Combo-2" antigen combinations of the invention). Individual symbols show the survival duration of individual mice; the horizontal bar for each group shows the median survival duration; the percentage figures are survival 14 days after challenge; and the p values at the top are t-Test comparisons of median survival durations between groups.

Figure 6 shows the number of colony forming units (cfu) in mouse kidneys after infection with 9×10^6 cfu of Newman strain in the abscess model. Horizontal bars are averages per group, and the figure beneath each group is the log reduction relative to the PBS control group.

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Figure 7 shows bacterial count (log CFU/ml) in kidneys of mice in an abscess model experiment. Mice were challenged with the following strains: (A) MW2; (B) LAC; (C) Staph19; or (D) MU50. Each point is an individual animal and the bar shows the median count per group. Mice had been immunised as shown on the x-axis label.

Figure 8 shows the formation of Sta011 oligomers in the presence of increasing concentrations of Ca⁺⁺ ions. Numbers indicate mM concentrations, and a * indicates the presence of 50mM EDTA.

Figure 9 shows IgG titers against (A) EsxAB (B) Sta006 (C) Hla-H35L (D) Sta011. Each graph has three groups, with a pair of bars per group. The right-hand bar in a pair shows pre-immune IgG and the left-hand bar shows post-immune IgG. The three groups are the compositions used for immunising and, from left to right, are: negative control of adjuvant alone; the Combo 1 combination; and the relevant antigen alone.

Figure 10 shows bacterial counts values (log CFU/ml) in mice after challenge with the indicated strains. Each point is an individual animal and the bar shows the median CFU. The P value beneath the IsdB and Combo columns is a comparison against the adjuvant-only control.

Figure 11 shows the area of abscesses (mm²) in mice after challenge with Newman strain.

Figure 12 shows days of survival of mice after challenge with four different strains: Newman (○), ST-80 (□), USA300-FPR3757 (Δ) or USA300-Lac (x) strains. Each point is an individual animal, the bar shows the median survival, and the heading number shows the % of animals surviving after 15 days. Mice received aluminium hydroxide adjuvant alone, IsdB or Combo1.

Figure 13 shows the median survival (days) of mice after challenge. The mice had been immunised with the antigens indicated on the X-axis. Each point is an individual animal and the bar shows the median survival. The heading numbers show the % of animals surviving after 15 days.

MODES FOR CARRYING OUT THE INVENTION

Antigen selection

[0595] *S. aureus* proteins have been selected for use as vaccine components based on various criteria.

[0596] IsdA is a surface protein involved in iron uptake. It is detectable with a high molecular weight (>250kDa) in immunoblots of whole cell lysates and cell wall fractions of *S. aureus*. Furthermore, labelled anti-IsdA antibodies revealed extracellular structures. These structures were seen in a variety of growth and infection conditions, including iron positive conditions (in which IsdA expression is reported to be suppressed). The structures have a tail up to 4μm long, with a typical orientation parallel to the mammalian cell surface. Detached IsdA-positive structures were observed to adhere on the surface of epithelial cells, but lose cell junction localization. Epithelia/bacteria interaction may stimulate expression of the structures. In addition, the inventors have found that IsdA is well conserved between different strains (present in 36/36 strains tested; see below), thus offering protection across a broad population of circulating strains. Iron uptake is important for virulence, so the protein is likely to be available for immune attack at pathological stages of the bacterial life cycle. The inventors have found that the protein is not cytotoxic to human cells (see below). The protein can also adsorb reasonably well to aluminium hydroxide (see below), which is useful for stable formulation for delivery to humans. It is useful for providing an immune response to prevent hematic spread of the bacterium.

[0597] EsxA and EsxB are small acidic dimeric secreted proteins. The inventors have found that EsxA is highly conserved between different strains (present in 36/36 strains tested; see below), while EsxB is present in 25/36 strains. The proteins are involved in persisting an infection and so are likely to be available for immune attack at pathological stages of the bacterial life cycle. The inventors have found that a fusion of EsxA and EsxB ('EsxAB') is not cytotoxic to human cells (see below). It can also adsorb well to aluminium hydroxide (see below), which is useful for stable formulation for delivery to humans. Thus the antigens are useful for providing an immune response to prevent hematic spread of the bacterium.

[0598] Hla is a pore-forming secreted toxin. This protein is well conserved between different strains (present in 36/36 strains tested; see below), thus offering protection across a broad population of circulating strains. It is an important virulence factor so is likely to be available for immune attack at pathological stages of the bacterial life cycle. It is not cytotoxic to human cells (see below). The protein can adsorb reasonably well to aluminium hydroxide (see below), which is useful for stable formulation for delivery to humans. It is useful for providing an immune response to prevent pneumonia.

[0599] Spa is a surface protein involved in Fc binding. The inventors have found that this protein is well conserved between different strains (present in 36/36 strains tested), thus offering protection across a broad population of circulating

strains. It is important for virulence so is likely to be available for immune attack at pathological stages of the bacterial life cycle. The protein can also adsorb reasonably well to aluminium hydroxide (see below), which is useful for stable formulation for delivery to humans. It is useful for providing an immune response to prevent hematic spread of the bacterium.

5 **[0600]** Sta006 (also known as FhuD2) is a surface protein involved in iron uptake. The inventors have found that this protein is well conserved between different strains (present in 36/36 strains tested; see below), thus offering protection across a broad population of circulating strains. The inventors have found that the protein is not cytotoxic to human cells (see below). The protein can also adsorb well to aluminium hydroxide (see below), which is useful for stable formulation for delivery to humans. It is useful for providing an immune response to prevent hematic spread of the bacterium.

10 **[0601]** Sta011 is a surface lipoprotein. The inventors have found that this protein is well conserved between different strains (present in 36/36 strains tested; see below), thus offering protection across a broad population of circulating strains. The inventors have found that the protein is not cytotoxic to human cells (see below). The protein can also adsorb reasonably well to aluminium hydroxide (see below), which is useful for stable formulation for delivery to humans. It is useful for providing an immune response to prevent hematic spread of the bacterium. This protein has been shown to assemble into oligomers in the presence of Ca⁺⁺ ions, but not Mg⁺⁺ ions (see Figure 8). These experiments used 5µg recombinant tag-free Sta011, incubated at 37°C for 25 minutes with increasing CaCl₂ concentrations from 0.5-50mM, then analysed by gel electrophoresis on a clear native gel. A mobility shift (indicating oligomerisation) was evident from 2mM Ca⁺⁺, and particularly >5mM. These levels compare to blood Ca⁺⁺ concentrations of about 1.2mM, serum concentrations of about 11mM, and milk concentrations of about 32mM. EDTA reversed the shift.

20 **[0602]** Surface digestion [302] and/or analysis of secreted proteins revealed peptide fragments from ClfA, ClfB, coA, eap, ebhA, ebpS, efb, emp, FnBA, FnBB, hla, lsdA, lsdB, lsdH, ukD, lukS, sdrD, sdrE, sasB, sasD, sasF, spa, sta001, sta002, sta003, sta004, sta005, sta006, sta007, sta008, sta009, sta010, sta011, sta019, sta023, sta024, sta028, sta036, sta040, sta049, sta050, sta054, sta057, sta064, sta065, sta073, sta095, sta096, sta098, sta100, sta101, sta102, sta103, sta105, sta107, sta108, sta109, sta111, sta112, sta113, sta115, sta116, sta117, sta118, sta120, NW_06, NW_07, NW_08, NW_09 and NW_10 e.g. SEQ ID NOs: 228 and 229 were identified as fragments of sta019.

25 **[0603]** Conjugated capsular saccharides are useful for providing opsonic immunity. Serotypes 5 and 8 cover about 85% of clinical isolates.

Strain coverage

30 **[0604]** A panel of 36 clinical isolates was used to represent circulating strains, including strains belonging to the five clonal lineages representing the vast majority of worldwide circulating CA-MRSA (community-associated methicillin-resistant *S. aureus*). HA-MRSA (hospital-associated MRSA) and non-MRSA strains were also included. Overall the panel included 9 HA-MRSA strains, 7 CA-MRSA strains, 2 MRSA strains, and 18 other strains.

35 **[0605]** Genes encoding lsdA, Hla, EsxA, Sta006, Sta011, Spa, and ClfB were present in all 36 strains. The gene for EsxB was absent from 11/36 strains, and the gene for SdrD was absent from 6/36 strains.

[0606] The encoded lsdA sequences were 95-100% identical across the panel, and the protein was expressed in iron-limited conditions in the stationary growth phase. The encoded SdrD sequences were 95-100% identical in the 30/36 SdrD^{+ve} panel members. The encoded EsxA sequences were 100% identical across the panel; the encoded EsxB sequences were 95-100% identical in the 25 EsxB^{+ve} strains. The encoded ClfB sequences were 93-100% identical across the panel, and this protein was also found to be highly surface-exposed in the early exponential growth phase.

[0607] Conservation in the encoded amino acid sequences were as follows (% identity):

Antigen	lsdA	ClfB	SdrD	Spa	Hla	EsxA	EsxB	Sta006
%	95-100	97-100	88-100	98-100	97-100	100	95-100	99.7-100

45 **[0608]** A larger panel of 61 strains was screened for the presence of genes encoding Hla and Sta006, as well as for their expression. This panel covered both MRSA and MSSA strains, a variety of geographical origins, and a variety of ST and clonal complex types. 9/61 strains did not express Hla, whereas all but one strain expressed Sta006 (data for the 61st strain were inconclusive). Thus a vaccine based on Hla alone is unlikely to give adequate coverage for a universal vaccine, but this problem could be overcome by addition of Sta006.

Cytotoxicity and cell binding studies

55 **[0609]** The analysis of the potential cellular cytotoxicity by *S. aureus* recombinant antigens Hla, Hla-H35L, lsdA, lsdB, sta006, sta011 and EsxAB was conducted on HBMECs and A549 cells. Annexin V and propidium iodide staining were

used to measure the percentage of early and late apoptotic cells by flow cytometry. Endothelial cells were grown in 24 well plates up to fully confluent. Cells were then incubated for 24 hours with three different concentration of recombinant antigens (10 μ g/ml, 1 μ g/ml, 0.1 μ g/ml). The combination of TNF- α and cycloheximide (CHX), which has been reported to induce apoptosis in endothelial cells, was used as a positive control. Incubation with PBS buffer alone was a negative control. Analysis was then performed by FACS.

[0610] None of the antigens induced a cytotoxic effect on HBMECs or A549 cells. Indeed, the percentage of live cell population compared to control cells remained essentially constant up to 24 hours of incubation. In contrast, the combination of TNF- α and CHX induced a 25% increase in the number of apoptotic cells.

[0611] HBMECs were also used as an *in vitro* model for testing the binding of *S.aureus* recombinant antigens to human endothelial cells. HBMECs were grown up to confluence at 37°C in humidified atmosphere in RPMI 1640 medium supplemented with 10% heat-inactivated fetal calf serum, 10% NuSerum, 2mM glutamine, 1 mM pyruvate, 1% non-essential amino acids, 1% MEM vitamins, 100 units/ml penicillin, and 100 μ g/ml streptomycin. Binding of recombinant antigens to the cells was tested by indirect immunofluorescence and analyzed by FACS. The cells positive for binding were measured as net mean intensity of fluorescence respect to negative controls, identified as unspecific antibody recognition. Binding experiments were performed at 4°C. Mouse polyclonal antibodies specific for each of the recombinant antigens were used as primary antibodies and binding was detected by R-Phycoerythrin-conjugated goat anti-mouse IgG secondary antibody. As negative control, HBMECs were incubated with primary polyclonal antibodies detected by fluorescence-labeled secondary antibody or fluorescence-labeled secondary antibody alone. Binding of a known surfaced-exposed GBS antigen was used as positive control.

[0612] Hla and Hla-H35L were the only antigens able to strongly bind to endothelial cells. The haemolytic activity of these two antigens was also tested.

[0613] De-fibrinated sheep and rabbit blood were used to measure their haemolytic activity by spectrophotometric assay. The blood was incubated at 37°C for 30 minutes with serial dilution 1:4 of the two proteins. Incubation with water, to cause osmotic lysis, and incubation with a *S.pyogenes* protein, were positive controls; as negative control, the blood was incubated with PBS+ BSA 0.5%.

[0614] Recombinant native Hla, but not its H35L mutant form, showed haemolytic activity on rabbit erythrocytes. The mutant was at least 150-fold less haemolytic than wild-type. Both proteins had no haemolytic activity on sheep blood.

[0615] Thus the *S.aureus* recombinant vaccine candidates do not show any cytotoxicity both on A549 epithelial cell line and HBMEC endothelial cell line. Importantly, Hla, a secreted toxin known to form pores into the plasma membrane of host cells, could bind A549 cells but did not induce cytotoxicity on them; it was also able to induce haemolysis of rabbit erythrocytes. In contrast, recombinant Hla-H35L, a variant toxin with a single amino acid substitution that cannot form cytolytic pores, did not induce cellular damage in both human cell lines and rabbit erythrocytes. These findings indicate that this mutant form of Hla may be more safely used in a vaccine composition. None of the other antigens showed the capacity to bind to host cells.

Adjuvant formulation

[0616] Selected *S.aureus* protein antigen candidates have been formulated with aluminium hydroxide, either individually or as a combination of proteins, with or without capsular polysaccharide conjugate(s). The formulations have been optimized for pH and osmolarity.

[0617] The antigens were EsxA-B, Sta006, Sta011, Hla-H35L, SdrD, IsdA, IsdA₄₀₋₁₈₄, Sta019, Sta021, Sta073, ClfB₄₅₋₅₅₂, SdrD₅₃₋₅₉₂, SasF, and IsdB. These are formulated as monovalent antigens at 100 μ g/ml, or as combinations at 50 μ g/ml each. Capsular saccharide conjugates from type 5 or type 8 strains are added at 5 μ g/ml, 10 μ g/ml or 25 μ g/ml. Aluminium hydroxide was used at 2mg/ml, in a 10mM histidine buffer (pH 6.5) and with 9mg/ml NaCl.

[0618] All monovalent and combination formulations, with or without conjugates, could be adjusted with respect to a desired pH and osmolality. The formulations had pH in the range 6.2-7.3, and osmolality in the range 248-360 mOsm/kg. Glycerol was excluded from formulations as it had a negative impact on osmolality.

[0619] All proteins tested, in various monovalent and combination formulations, adsorbed well to the aluminium hydroxide adjuvant, except for IsdA, IsdA₄₀₋₁₈₄, Sta019, and Sta073.

[0620] The individual Sta006, Sta011, EsxA-B and Hla-H35L proteins were completely adsorbed, and could be desorbed without altering their pre-adsorption electrophoretic profile.

[0621] Each antigen in a combination of Sta006, Sta011, EsxA-B and Hla-H35L was completely adsorbed, with no inter-antigen competition for the adjuvant. The antigens in a combination of Sta006, Sta011, EsxA-B and IsdA₄₀₋₁₈₄ were also completely adsorbed, except for IsdA₄₀₋₁₈₄, which behaved in the same way as the monovalent protein. For both combinations, the antigens could be desorbed without altering their pre-adsorption electrophoretic profile.

[0622] The additional presence of type 5 and/or type 8 conjugates also did not change the adsorption or desorption characteristics of the antigens e.g. in combination with Sta006+Sta011+EsxA-B.

[0623] A short stability study (2 weeks at 4°C) was performed to evaluate the stability of monovalent formulations and

to evaluate antigen integrity. All tested formulations were stable for their pH and osmolality. All antigens remained completely adsorbed to the adjuvant. All antigens maintained their desorption characteristics. There was no evidence of increased degradation or aggregation of antigens after desorption.

5 Efficacy testing

[0624] Individual antigens sta006, sta011, sta012, sta017, sta019, sta021 and sta028 were tested for their ability to protect against IV challenge by 1.2×10^7 cfu of Newman strain (type 5). Results are shown in Figure 1. All antigens reduced bacterial numbers compared with the control, and the best results were seen with sta006, sta011 and sta019.

[0625] Further individual antigens were tested: (i) NW_10; (ii) IsdA_{40-184} ; (iii) Sta002; (iv) Sta003; (v) Sta073; (vi) Sta101; (vii) Sta014; (viii) Hla-PSGS; (ix) $\text{SdrD}_{\text{CnaB}}$. The increase in survival, compared to the negative control group, 15 days after challenge was: (i) 50%; (ii) 19%; (iii) 37%; (iv) 43%; (v) 25%; (vi) 12%; (vii) 25%; (viii) 56%; (ix) 39%.

[0626] Two hybrid polypeptides were also tested: (i) HlaH35L-EsxAB; (ii) Sta006-EsxAB. The increase in survival after challenge, compared to the negative control group, was: (i) 25%; (ii) 25%.

[0627] Table 2 gives a summary of results obtained with various antigens in the abscess model.

[0628] Experiment SA-10-a tested the efficacy of antigen combinations. Six groups of twelve CD-1 mice received a negative control (PBS), IsdB , or one of the following combinations, adjuvanted with aluminium hydroxide: (i) EsxAB + Hla-H35L; (ii) Sta006 + Sta011 + EsxAB; (iii) Sta006 + Sta011 + EsxAB + Hla-H35L; or (iv) Sta006 + Sta011 + IsdA_{40-184} + EsxAB. Two administrations were given, at days 0 and 14. At day 24 mice received 3×10^8 cfu of Newman strain staphylococcus and survival in each group was assessed every 24 hours for two weeks. Results are shown in Figure 2. After 14 days, 25% of animals in the positive control group had survived, but 50% of animals in group (ii) had survived, as had 58% of animals in groups (iii) & (iv), and 75% in group (i).

[0629] Experiment SA-10-b used the same methods to test: (i) ClfB_{45-552} + Hla-H35L + Sta006 + EsxAB; (ii) ClfB_{45-552} + Sta011 + Sta006 + EsxAB; (iii) ClfB_{45-552} + IsdA_{40-184} + Sta006 + EsxAB; or (iv) SdrD_{53-592} + IsdA_{40-184} + Sta006 + EsxAB. Results are shown in Figure 3. After 14 days, 25% of animals in the positive control group and in group (ii) had survived, but 33% of animals in group (iv) had survived, 75% of animals in group (i), and 83% of animals in group (iii).

[0630] Further combinations were also used to immunise mice. The combinations were typically adjuvanted with aluminium hydroxide (see above) and were administered on days 0 and 14. The immunisations were in CD1 mice, 12 per group. On day 24 the mice were challenged with a lethal dose of live bacteria and survival was then followed for 14 further days. For comparison, PBS was used as a negative control and IsdB as a positive control [2].

[0631] Experiment SA-11 tested: (i) a type 5 conjugate combined with EsxAB + Sta006 + Sta011; (ii) EsxAB + Sta019 + Sta006 + Sta011; (iii) a type 5 conjugate + Hla-H35L + Sta006 + Sta011; (iv) EsxAB + Hla-H35L + Sta006 + Sta011; or (v) EsxAB + IsdA_{40-184} + Sta006 + Sta011. 14 days after challenge all of the negative control animals had died, but 42% of positive control animals had survived. Survival results in the test groups were as follows: (i) 67%; (ii) 42%; (iii) 75%; (iv) 33%; and (v) 25%.

[0632] Experiment SA-12 tested: (i) Hla-H35L + IsdA_{40-184} + Sta006 + Sta011; (ii) Hla-H35L + EsxAB + Sta006 + Sta011; (iii) EsxAB + IsdA_{40-184} + Sta006 + Sta011; (iv) EsxAB + IsdA + Sta006 + Sta011. 14 days after challenge 8% of the negative control animals and 17% of positive control animals had survived. Survival results in the test groups were as follows: (i) 50%; (ii) 50%; (iii) 25%; (iv) 33%.

[0633] Experiment SA-14 tested: (i) EsxAB + Hla-H35L + Sta006 + Sta011; (ii) EsxAB + IsdA_{40-184} + Sta006 + Sta011; (iii) Sta006 + Sta011 + Sta019 + EsxAB; (iv) Sta006 + Sta011 + Sta019 + Hla-H35L. 14 days after challenge with 5×10^8 CFU of Newman strain, 18% of the negative control animals and 9% of positive control animals had survived; survival results in the test groups were as follows: (i) 58%; (ii) 67%; (iii) 42%; (iv) 83%. Survival numbers over 14 days are shown in Figure 4, showing that all combinations performed better than the two controls on every post-challenge day.

[0634] Experiment SA-17a tested: (i) EsxAB + Sta006 + Sta011 + serotype 5 conjugate + serotype 8 conjugate; (ii) EsxAB + Sta073 + Sta011 + serotype 5 conjugate + serotype 8 conjugate; (iii) EsxAB + Hla-H35L + Sta011 + Sta073. Compared to the negative control, the increase in survival 15 days after challenge with Newman strain was: (i) 17%; (ii) 42%; (iii) 34%. The median survival in groups (ii) and (iii) was the full 15 days, and was 12 days in group (i).

[0635] Further antigen combination experiments tested: (a) serotype 5 conjugate + serotype 8 conjugate + EsxAB + Sta006 + Sta011; (b) Sta002+Sta003+Sta021+NW-10; (c) EsxAB+ HlaH35L + Sta006 + Sta019; and (d) EsxAB + Sta006+Sta019. Compared to the negative control, the increase in survival after challenge with Newman strain was: (a) 37%; (b) 36%; (c) 13%; and (d) 0%.

[0636] Survival data from studies SA-10, SA-11, SA-12 and SA-14 were combined to assess the efficacy of two combinations when compared to PBS or IsdB . "Combo-1" was EsxAB+Hla-H35L+Sta006+Sta011 (with polypeptides comprising SEQ ID NOs: 241, 150, 246 & 247). "Combo-2" was EsxAB+ IsdA_{40-184} +Sta006+Sta011. The median survival times for each group of 48 mice after 14 days were compared. Whereas the PBS and IsdB groups had a median survival time of 1 day, mice in the "Combo-1" and "Combo-2" groups had a median survival time of 14 days. The differences in median survival duration were compared by a t-test: survival in the "Combo-1" group was statistically superior to both

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the PBS group ($p < 0.0001$) and the lsdB group ($p < 0.0001$); survival in the "Combo-2" group was statistically superior to both the PBS group ($p < 0.0001$) and the lsdB group ($p = 0.0049$). These data are shown in Figure 5.

[0637] Figure 6 shows data with Combo-1 and Combo-2 in the abscess model. Kidneys of mice are isolated after challenge and are then homogenised and plated. The cfu count indicates the level of abscess formation. Figure 6 shows data from a single experiment. The numbers beneath the data show the log reduction relative to the PBS group. The reduction is bigger in the two combination groups than with lsdB alone, with U-test (one tail) values of 0.0001 for Combo-1 and 0.0005 for Combo-2. The same effect was seen in the two combination groups in a second experiment in which an lsdB group was not included.

[0638] Further experiments compared protection achieved with Combo-1, lsdB or PBS against challenge with three different strains: Staph-19, FPR3757(USA300) and Lac(USA300). There were 44 mice per group and results were as follows (see also Figure 12), including one-tailed p-values for the survival proportion, where: P1 compares Combo-1 with PBS; P2 compares Combo-1 with lsdB; and P3 compared PBS with lsdB:

	Staph-19		FPR3757		Lac	
Survival	%	Days	%	Days	%	Days
PBS	20	1	45	8	47	7
lsdB	32	1	52	15	61	15
Combo-1	80	15	91	15	89	15
P1	<0.0001	-	<0.0001	-	0.0001	-
P2	<0.0001	-	<0.0004	-	0.0052	-
P3	0.1715	-	0.2137	-	0.1789	-

[0639] Further experiments showed that immunisation with adjuvanted Combo 1 reduced CFU counts after challenge with Newman, USA100, CC30 and USA300 strains, when compared to immunisation with adjuvant alone (aluminium hydroxide) or lsdB. Figure 10 shows CFU values (log/ml) for the four challenge strains. The lowest count, with $p < 0.015$ in each case, was achieved with Combo 1. The area of abscess was also assessed and was also lower in the Combo1-immunised mice (e.g. Figure 11).

[0640] Further experiments showed that Combo1 is highly protective against clinically relevant strains in the sepsis model, and always achieved a higher survival % than lsdB. Figure 12 shows that the median survival in Combo1-immunised mice (40 per group, 3 experiments) was the full 15 days when challenged with Newman, ST-80, FPR3757 or Lac strains, and that the proportion of mice surviving was $\geq 75\%$. In contrast, the median survival in lsdB-immunised mice was only 1 day with Newman and ST-80 challenge, with $< 65\%$ survival for all four challenge strains.

Comparison of Combo1 to its individual polypeptides

[0641] Various tests were performed to compare Combo1 to its four individual polypeptides (*i.e.* EsxA-B, Hla-H35L, Sta006, Sta011), as well as to lsdB or to an antigen-free negative control.

[0642] The opsonophagocytic activity of sera from immunised animals was tested. Sera were obtained using (i) the four individual polypeptides, (ii) all pairs of the polypeptides, (iii) all triplets, or (iv) the full Combo 1 combination. For comparison, anti-lsdB serum was used. Pre-immune and negative control sera showed no killing of Newman strain in this assay. In a first experiment: anti-lsdB serum showed 27% killing; sera against each of the four individual polypeptides showed between 26-34% killing; all multi-polypeptide combinations showed at least 34% killing; and sera raised with Combo-1 showed 39% killing. In a second experiment sera with Combo-1 showed 43% killing but anti-lsdB serum performed slightly better; all single or multi-polypeptide sera using the Combo-1 polypeptides showed at least 26% killing.

[0643] Further experiments looked at passive protection achieved by transferring into mice (20 per group, 8 week old CD1 mice) antiserum from immunised rabbits. Four groups received 200 μ l of sera from rabbits immunised with one of EsxA-B, Hla-H35L, Sta006, Sta011; a fifth group received 50 μ l of each serum (200 μ l in total). Two other groups received serum from lsdB-immunised rabbits or serum from rabbits immunised with saline+adjuvant. 15 minutes later the mice were challenged intraperitoneally (10^8 CFU of Newman strain) and then mortality was assessed after 14 days. Results were as follows:

	EsxA-B	Sta006	Sta011	HlaH35L	Combo1	lsdB	-ve ctrl
Survival	5%	26%	0%	15%	25%	10%	5%

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[0644] In further experiments the level of specific antibodies induced in CD1 mice were examined to assess the immunogenicity of the four polypeptides in Combo1. Compositions included either 20 μ g of each of the four single polypeptides, or 4x10 μ g in the combination. The compositions included an aluminium hydroxide adjuvant. Serum levels of antigen-specific IgG were determined by Luminex 4Plex assay As shown in Figure 9, all four polypeptides were highly immunogenic in CD1 mice on their own and in combination. In each case the titer against a polypeptide was higher when it was administered in the combination than when administered alone (compare middle and right pairs).

[0645] Further experiments compared protection achieved either with Combo-1 or with its four individual polypeptides. IsdB was also included for comparison. The proportions of animals surviving (40 animals per group) 15 days after challenge with Newman strain, and the average (median) survival in days, were as follows, including a one-tailed p-value of the surviving proportion in comparison with a PBS+adjuvant negative control:

	EsxA-B	Sta006	Sta011	HlaH35L	Combo1	IsdB	PBS
Survival	34%	28%	16%	39%	59%	22%	5%
p	0.0017	0.0003	0.0064	<0.0001	<0.0001	0.0006	-
Days	1	2	1	10	15	1	0

[0646] The murine abscess model was used to compare the four individual polypeptides with the Combo 1 combination. In some experiments mice were immunised with IsdB for comparison. Antigens were adjuvanted with aluminium hydroxide, and adjuvant alone was used as a negative control. Figure 7 shows the numbers of bacteria in animals' kidneys after challenge with four different strains. The lowest average counts were seen for the Combo 1 combination.

[0647] Challenge experiments were performed following immunisation with (i) the four individual polypeptides, (ii) all pairs, (iii) all triplets, or (iv) the full Combo1 combination. IsdB or buffer alone were used for comparison. Survival results from 24 mice per group (3 experiments) after challenge with 5x10⁸ CFU of Newman strain are shown in Figure 13. The median survival for IsdB was only 2 days. The median survival for the individual Combo1 polypeptides ranged from 1-6 days. Pairs of the polypeptides gave median survival of 2-11 days. Triplets gave median survival of 8-15 days. The full Combo1 combination gave a median survival of the full 15 days, with 59% of mice surviving this long (*cf.* only 35% with IsdB).

[0648] It will be understood that the invention has been described by way of example only and modifications may be made whilst remaining within the scope and spirit of the invention.

TABLE 1: NOMENCLATURE CROSS-REFERENCE

SEQ ID NO	Name	NCTC 8325 strain		Newman strain	
		SAOUHSC_#	GI	NWMN_#	GI
1	clfA	SAOUHSC_00812	88194572	NWMN_0756	151220968
2	clfB	SAOUHSC_02963	88196585	NWMN_2529	151222741
3	coA	SAOUHSC_00192	88194002	NWMN_0166	151220378
4	eap	SAOUHSC_02161	88195840	NWMN_1872	151222084
5	ebhA	SAOUHSC_01447	88195168	-	-
6	ebpS	SAOUHSC_01501	88195217	NWMN_1389	151221601
7	efb	SAOUHSC_01114	88194860	NWMN_1069	151221281
8	emp	SAOUHSC_00816	88194575	NWMN_0758	151220970
9	esaC	SAOUHSC_00264	88194069	-	-
10	esxA	SAOUHSC_00257	88194063	-	-
11	esxB	SAOUHSC_00265	88194070	-	-
12	FnBA	SAOUHSC_02803	88196438	NWMN_2399	151222611
13	FnBB	SAOUHSC_02802	88196437	NWMN_2397	151222609
14	hla	SAOUHSC_01121	88194865	NWMN_1073	151221285
15	hlgB	SAOUHSC_02710	88196350	-	-

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SEQ ID NO	Name	NCTC 8325 strain		Newman strain	
		SAOUHSC_#	GI	NWMN_#	GI
16	hlgC	SAOUHSC_02709	88196349	-	-
17	isdA	SAOUHSC_01081	88194829	NWMN_1041	151221253
18	isdB	SAOUHSC_01079	88194828	-	-
19	isdC	SAOUHSC_01082	88194830	-	-
20	isdG	SAOUHSC_01089	88194836	-	-
21	isdH	SAOUHSC_01843	88195542	NWMN_1624	151221836
22	isdI	SAOUHSC_00130	88193943	-	-
23	lukD	SAOUHSC_01954	88195647	NWMN_1718	151221930
24	lukE	SAOUHSC_01955	88195648	-	-
25	lukF	SAOUHSC_02241	88195914	-	-
26	lukS	SAOUHSC_02243	88195915	NWMN_1928	151222140
27	nuc	SAOUHSC_01316	88195046	-	-
28	sasA	SAOUHSC_02990	88196609	-	-
29	sasB	SAOUHSC_02404	88196065	-	-
30	sasC	SAOUHSC_01873	88195570	-	-
31	sasD	SAOUHSC_00094	88193909	-	-
32	sasF	SAOUHSC_02982	88196601	-	-
33	sdrC	SAOUHSC_00544	88194324	-	-
34	sdrD	SAOUHSC_00545	88194325	-	-
35	sdrE2	-	-	NWMN_0525	151220737
36	spa	SAOUHSC_00069	88193885	NWMN_0055	151220267
37	sta001	SAOUHSC_00025	88193846	NWMN_0022	151220234
38	sta002	SAOUHSC_00356	88194155	NWMN_0364	151220576
39	sta003	SAOUHSC_00400	88194195	NWMN_0401	151220613
40	sta004	SAOUHSC_00749	88194514	NWMN_0705	151220917
41	sta005	SAOUHSC_01127	88194870	NWMN_1077	151221289
42	sta006	SAOUHSC_02554	88196199	NWMN_2185	151222397
43	sta007	SAOUHSC_02571	88196215	NWMN_2199	151222411
44	sta008	SAOUHSC_02650	88196290	NWMN_2270	151222482
45	sta009	SAOUHSC_02706	88196346	NWMN_2317	151222529
46	sta010	SAOUHSC_02887	88196515	NWMN_2469	151222681
47	sta011	SAOUHSC_00052	88193872	-	-
48	sta012	SAOUHSC_00106	88193919	-	-
49	sta013	SAOUHSC_00107	88193920	-	-
50	sta014	SAOUHSC_00137	88193950	-	-
51	sta015	SAOUHSC_00170	88193980	-	-
52	sta016	SAOUHSC_00171	88193981	-	-

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SEQ ID NO	Name	NCTC 8325 strain		Newman strain	
		SAOUHSC_#	GI	NWMN_#	GI
53	sta017	SAOUHSC_00186	88193996	-	-
54	sta018	SAOUHSC_00201	88194011	-	-
55	sta019	SAOUHSC_00248	88194055	NWMN_0210	151220422
56	sta020	SAOUHSC_00253	88194059	-	-
57	sta021	SAOUHSC_00256	88194062	-	-
58	sta022	SAOUHSC_00279	88194083	-	-
59	sta023	SAOUHSC_00284	88194087	-	-
60	sta024	SAOUHSC_00300	88194101	-	-
61	sta025	SAOUHSC_00362	88194160	-	-
62	sta026	SAOUHSC_00404	88194198	-	-
63	sta027	SAOUHSC_00661	88194426	-	-
64	sta028	SAOUHSC_00671	88194436	NWMN_0634	151220846
65	sta029	SAOUHSC_00754	88194518	-	-
66	sta030	SAOUHSC_00808	88194568	-	-
67	sta031	SAOUHSC_00860	88194617	-	-
68	sta032	SAOUHSC_00958	88194715	-	-
69	sta033	SAOUHSC_00987	88194744	-	-
70	sta034	SAOUHSC_00988	88194745	-	-
71	sta035	SAOUHSC_00998	88194754	-	-
72	sta036	SAOUHSC_01084	88194831	-	-
73	sta037	SAOUHSC_01085	88194832	-	-
74	sta038	SAOUHSC_01088	88194835	-	-
75	sta039	SAOUHSC_01124	88194868	-	-
76	sta040	SAOUHSC_01125	88194869	NWMN_1076	151221288
77	sta041	SAOUHSC_01175	88194914	-	-
78	sta042	SAOUHSC_01180	88194919	-	-
79	sta043	SAOUHSC_01219	88194955	-	-
80	sta044	SAOUHSC_01508	88195223	-	-
81	sta045	SAOUHSC_01627	88195337	-	-
82	sta046	SAOUHSC_01918	88195613	-	-
83	sta047	SAOUHSC_01920	88195615	-	-
84	sta048	SAOUHSC_01949	88195642	-	-
85	sta049	SAOUHSC_01972	88195663	NWMN_1733	151221945
86	sta050	SAOUHSC_02127	88195808	-	-
87	sta051	SAOUHSC_02147	88195827	-	-
88	sta052	SAOUHSC_02246	88195918	-	-
89	sta053	SAOUHSC_02257	88195928	-	-

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SEQ ID NO	Name	NCTC 8325 strain		Newman strain	
		SAOUHSC_#	GI	NMWN_#	GI
90	sta054	SAOUHSC_02333	88195999	-	-
91	sta055	SAOUHSC_02448	88196100	-	-
92	sta056	SAOUHSC_02463	88196115	-	-
93	sta057	SAOUHSC_02576	88196220	NWMN_2203	151222415
94	sta058	SAOUHSC_02690	88196330	-	-
95	sta059	SAOUHSC_02708	88196348	-	-
96	sta060	SAOUHSC_02767	88196403	-	-
97	sta061	SAOUHSC_02783	88196419	-	-
98	sta062	SAOUHSC_02788	88196424	-	-
99	sta063	SAOUHSC_02971	88196592	-	-
100	sta064	SAOUHSC_03006	88196625	NWMN_2569	151222781
101	sta065	SAOUHSC_00051	88193871	-	-
102	sta066	SAOUHSC_00172	88193982	-	-
103	sta067	SAOUHSC_00176	88193986	-	-
104	sta068	SAOUHSC_00327	88194127	-	-
105	sta069	SAOUHSC_00427	88194219	-	-
106	sta070	SAOUHSC_00773	88194535	-	-
107	sta071	SAOUHSC_00854	88194612	-	-
108	sta072	SAOUHSC_00872	88194629	-	-
109	sta073	SAOUHSC_00994	88194750	NWMN_0922	151221134
110	sta074	SAOUHSC_01220	88194956	-	-
111	sta075	SAOUHSC_01256	88194989	-	-
112	sta076	SAOUHSC_01263	88194996	-	-
113	sta077	SAOUHSC_01317	88195047	-	-
114	sta078	SAOUHSC_01857	88195555	-	-
115	sta079	SAOUHSC_01935	88195630	-	-
116	sta080	SAOUHSC_01936	88195631	-	-
170	sta081	SAOUHSC_01938	88195633		
117	sta082	SAOUHSC_01939	88195634	-	-
118	sta083	SAOUHSC_01941	88195635	-	-
119	sta084	SAOUHSC_01942	88195636	-	-
120	sta085	SAOUHSC_02171	88195848	-	-
121	sta086	SAOUHSC_02327	88195993	-	-
122	sta087	SAOUHSC_02635	88196276	-	-
123	sta088	SAOUHSC_02844	88196477	-	-
124	sta089	SAOUHSC_02855	88196486	-	-
125	sta090	SAOUHSC_02883	88196512	-	-

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SEQ ID NO	Name	NCTC 8325 strain		Newman strain	
		SAOUHSC_#	GI	NWMN_#	GI
126	sta091	SAOUHSC_00685	88194450	-	-
127	sta092	SAOUHSC_00174	88193984	-	-
128	sta093	SAOUHSC_01854	88195552	-	-
129	sta094	SAOUHSC_01512	88195226	-	-
130	sta095	SAOUHSC_00383	88194180	NWMN_0388	151220600
131	sta096	SAOUHSC_00384	88194181	-	-
132	sta097	SAOUHSC_00386	88194182	-	-
133	sta098	SAOUHSC_00389	88194184	NWMN_0391	151220603
134	sta099	SAOUHSC_00390	88194185	-	-
135	sta100	SAOUHSC_00391	88194186	-	-
136	sta101	SAOUHSC_00392	88194187	NWMN_0394	151220606
137	sta102	SAOUHSC_00393	88194188	-	-
138	sta103	SAOUHSC_00394	88194189	-	-
139	sta104	SAOUHSC_00395	88194190	-	-
140	sta105	SAOUHSC_00399	88194194	NWMN_0400	151220612
141	sta106	SAOUHSC_01115	88194861	-	-
177	sta107	SAOUHSC_00354	88194153	NWMN_0362	151220574
178	sta108	SAOUHSC_00717	88194482	NWMN_0677	151220889
179	sta109	SAOUHSC_02979	88196599	NWMN_2543	151222755
180	sta110	SAOUHSC_01039	88194791		
181	sta111	SAOUHSC_01005	88194760	NWMN_0931	151221143
182	sta112	SAOUHSC_00634	88194402	NWMN_0601	151220813
183	sta113	SAOUHSC_00728	88194493	NWMN_0687	151220899
184	sta114	SAOUHSC_00810	88194570		
185	sta115	SAOUHSC_00817	88194576	NWMN_0759	151220971
186	sta116	SAOUHSC_01112	88194858	NWMN_1067	151221279
187	sta117	SAOUHSC_02240	88195913	NWMN_1926	151222138
188	sta118	SAOUHSC_01150	88194892	NWMN_1096	151221308
200	sta119	SAOUHSC_01100	88194846		
201	sta120	SAOUHSC_00365	88194163		
142	NW_6	-	-	NWMN_0757	151220969
143	NW_9	-	-	NWMN_0958	151221170
144	NW_10	-	-	NWMN_1066	151221278
145	NW_7	-	-	NWMN_1876	151222088
146	NW_8	-	-	NWMN_1877	151222089
147	NW_2	-	-	NWMN_1883	151222095
148	NW_1	-	-	NWMN_1924	151222136

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		NCTC 8325 strain		Newman strain	
SEQ ID NO	Name	SAOUHSC_#	GI	NMWN_#	GI
149	NW_5	-	-	NWMN_2392	151222604

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TABLE 2: ABSCESS MODEL RESULTS SUMMARY

	Immunising antigen(s)	Adjuvant	infecting strain & dose		Reduction**
5	Fnb	alum *	Newman	1.4E+07	2.13
	Sta005	alum	Newman	1.4E+07	1.26
	LukE	alum	Newman	1.4E+07	1.68
	SasD	alum	Newman	1.4E+07	0.10
10	SpA	alum	Newman	1.4E+07	0.41
	SasFHis	alum	Newman	1.4E+07	1.33
	CoA	alum	Newman	1.4E+07	1.01
	Sta028	alum	Newman	1.2E+07	1.85
	Sta017	alum	Newman	1.2E+07	1.23
15	Sta006	alum	Newman	1.2E+07	2.33
	Sta012	alum	Newman	1.2E+07	1.69
	Sta011	alum	Newman	1.2E+07	2.66
	Sta019	alum	Newman	1.2E+07	2.36
	Sta021	alum	Newman	1.2E+07	1.58
20	IsdA + EsxAB	alum	Newman	1.8E+07	0.11
	EsxAB	alum	Newman	1.8E+07	1.31
	NW_1	alum	Newman	1.8E+07	1.00
	NW_10	alum	Newman	1.8E+07	-0.65
25	Sta073	alum	Newman	1.8E+07	1.46
	Sta002	alum	Newman	1.8E+07	0.17
	Sta064	alum	Newman	1.8E+07	1.04
	Sta014	alum	Newman	1.8E+07	1.74
	Sta002	alum	Newman	1.0E+07	0.52
30	Sta014	alum	Newman	1.0E+07	1.02
	Sta064	alum	Newman	1.0E+07	1.22
	Sta006	alum	Newman	1.0E+07	0.80
	Sta073	alum	Newman	1.0E+07	0.92
	NW_1	alum	Newman	1.0E+07	0.77
35	NW_10	alum	Newman	1.0E+07	2.25
	Sta017	alum	Newman	1.0E+07	2.13
	Sta028	alum	Newman	1.0E+07	0.64
	Sta021	alum	Newman	1.0E+07	1.03
	Sta019	alum	Newman	1.0E+07	1.28
40	Sta011	alum	Newman	1.0E+07	0.78
	IsdB	alum	Newman	1.0E+07	1.22
	IsdA ₄₀₋₁₈₄	none	Newman	1.0E+07	0.58
	Sta006	none	Newman	1.0E+07	0.30
	Sta011	none	Newman	1.0E+07	0.62
45	EsxAB	none	Newman	1.0E+07	1.09
	Sasf	none	Newman	1.0E+07	0.11
	IsdB	none	Newman	1.0E+07	0.93
	IsdA ₄₀₋₁₈₄	alum	Newman	1.0E+07	1.02
	Sta006	alum	Newman	1.0E+07	0.45
50	Sta011	alum	Newman	1.0E+07	0.80
	EsxAB	alum	Newman	1.0E+07	0.47
	Sasf	alum	Newman	1.0E+07	-0.78
	IsdB	alum	Newman	1.0E+07	1.24
	Type 5 conjugate + IsdA ₄₀₋₁₈₄	alum	Newman	1.5E+07	0.34
55	Type 5 conjugate	alum	Newman	1.5E+07	0.72
	IsdA ₄₀₋₁₈₄	alum	Newman	1.5E+07	1.08

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	Type 5 conjugate	MF59	Newman	1.5E+07	0.45
	IsdB	alum	Newman	1.5E+07	1.50
	ClfB ₄₅₋₅₅₂	alum	Newman	1.5E+07	-0.05
5	Sta019	alum	Newman	1.5E+07	0.82
	IsdA ₄₀₋₁₈₄ + ClfB ₄₅₋₅₅₂	alum	Newman	1.5E+07	0.72
	Type 8 conjugate	alum	Becker	4.0E+07	1.51
	Type 8 conjugate	MF59	Becker	4.0E+07	0.35
10	EsxAB + Sta019 + Sta006 + Sta011	alum	Newman	1.0E+07	1.54
	combol	alum	Newman	1.0E+07	2.04
	EsxAB + IsdA ₄₀₋₁₈₄ + Sta006 + Sta011	alum	Newman	1.0E+07	0.84
	SdrD ₅₃₋₅₉₂	alum	Newman	1.0E+07	1.15
	Sta105	alum	Newman	1.0E+07	0.54
15	Sta101	alum	Newman	1.0E+07	1.51
	Sta116	alum	Newman	1.0E+07	1.23
	Sta106	alum	Newman	1.0E+07	1.20
	Sta107	alum	Newman	1.0E+07	1.77
	Sta004	alum	Newman	1.0E+07	0.70
20	Sta003	alum	Newman	1.0E+07	1.32
	EsxAB + Sta019 + Sta006 + Sta011	alum	Newman	9.0E+06	3.04
	combol	alum	Newman	9.0E+06	2.53
	EsxAB + IsdA ₄₀₋₁₈₄ + Sta006 + Sta011	alum	Newman	9.0E+06	1.85
	SdrD ₅₃₋₅₉₂	alum	Newman	9.0E+06	1.80
25	Sta105	alum	Newman	9.0E+06	0.60
	Sta101	alum	Newman	9.0E+06	0.83
	Sta116	alum	Newman	9.0E+06	1.96
	Sta106	alum	Newman	9.0E+06	2.56
	IsdB	alum	Newman	9.0E+06	1.37
30	Sta004	alum	Newman	9.0E+06	1.01
	Sta003	alum	Newman	9.0E+06	2.20
	IsdB	alum	Newman	1.0E+07	0.83
	Sta107	alum	Newman	1.0E+07	0.24
35	SrdC ₅₁₋₅₁₈	alum	Newman	1.0E+07	0.84
	SdrE ₅₃₋₆₃₂	alum	Newman	1.0E+07	1.08
	Hla ₂₇₋₇₆	alum	Newman	1.0E+07	0.18
	EsxAB + HlaH35L + Sta006 + Sta021	alum	Newman	1.0E+07	0.59
	EsxAB + HlaH35L + Sta006 + Sta019	alum	Newman	1.0E+07	0.85
40	EsxAB + HlaH35L + Sta006 + Sta017	alum	Newman	1.0E+07	1.88
	EsxAB + Hla ₂₇₋₇₆ + Sta006 + Sta021	alum	Newman	1.0E+07	1.49
	Hla ₂₇₋₇₆ + Sta006 + Sta017 + Sta019	alum	Newman	1.0E+07	0.00
	IsdB	alum	Newman	1.2E+07	1.07
	Sta107	alum	Newman	1.2E+07	1.35
45	SrdC ₅₁₋₅₁₈	alum	Newman	1.2E+07	2.17
	SdrE ₅₃₋₆₃₂	alum	Newman	1.2E+07	2.82
	Hla ₂₇₋₇₆	alum	Newman	1.2E+07	0.17
	EsxAB + HlaH35L + Sta006 + Sta021	alum	Newman	1.2E+07	1.70
	EsxAB + HlaH35L + Sta006 + Sta019	alum	Newman	1.2E+07	1.20
50	EsxAB + HlaH35L + Sta006 + Sta017	alum	Newman	1.2E+07	1.52
	EsxAB + Hla ₂₇₋₇₆ + Sta006 + Sta021	alum	Newman	1.2E+07	1.81
	Hla ₂₇₋₇₆ + Sta006 + Sta017 + Sta019	alum	Newman	1.2E+07	0.89
	IsdB	alum	Mu-50	3.8E+07	0.44
	Combol	alum	Mu-50	3.8E+07	1.73
55	IsdB	alum	USA 200	2.0E+07	1.17
	Combol	alum	USA 200	2.0E+07	1.87

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	IsdB	alum	USA 300	3.0E+07	0.09
	Combo1	alum	USA 300	3.0E+07	2.19
	IsdB	alum	Staph19	2.7E+07	0.66
5	Combo1	alum	Staph19	2.7E+07	0.46
	IsdB	alum	Mu-50	4.5E+07	0.98
	Combo1	alum	Mu-50	4.5E+07	0.76
	IsdB	alum	USA 200	1.6E+07	0.18
10	Combo1	alum	USA 200	1.6E+07	0.19
	IsdB	alum	USA 300	2.2E+07	-0.21
	Combo1	alum	USA 300	2.2E+07	-0.29
	IsdB	alum	Staph19	2.3E+07	0.57
	Combo1	alum	Staph19	2.3E+07	0.80
15	IsdB	alum	LAC	3.50E+07	2.67
	Sta011	alum	LAC	3.50E+07	1.35
	EsxAB	alum	LAC	3.50E+07	2.21
	HlaH35L	alum	LAC	3.50E+07	0.71
	Sta006	alum	LAC	3.50E+07	2.39
20	Combo1	alum	LAC	3.50E+07	2.66
	IsdB	alum	MW2	3.00E+07	1.17
	Sta011	alum	MW2	3.00E+07	0.82
	EsxAB	alum	MW2	3.00E+07	1.39
	HlaH35L	alum	MW2	3.00E+07	0.87
25	Sta006	alum	MW2	3.00E+07	0.91
	Combo1	alum	MW2	3.00E+07	2.69
	IsdB	alum	LAC	4.00E+07	1.54
	Sta011	alum	LAC	4.00E+07	1.95
	EsxAB	alum	LAC	4.00E+07	1.31
30	HlaH35L	alum	LAC	4.00E+07	0.75
	Sta006	alum	LAC	4.00E+07	1.74
	Combo1	alum	LAC	4.00E+07	2.21
	IsdB	alum	MW2	2.75E+07	1.22
35	Sta011	alum	MW2	2.75E+07	1.25
	EsxAB	alum	MW2	2.75E+07	1.16
	HlaH35L	alum	MW2	2.75E+07	1.61
	Sta006	alum	MW2	2.75E+07	1.13
	Combo1	alum	MW2	2.75E+07	1.97
40	Sta011	alum	Mu-50	4.00E+07	1.10
	EsxAB	alum	Mu-50	4.00E+07	0.86
	HlaH35L	alum	Mu-50	4.00E+07	0.71
	Sta006	alum	Mu-50	4.00E+07	1.57
	Combo1	alum	Mu-50	4.00E+07	1.72
45	Sta011	alum	Staph19	5.30E+07	1.23
	EsxAB	alum	Staph19	5.30E+07	1.19
	HlaH35L	alum	Staph19	5.30E+07	0.65
	Sta006	alum	Staph19	5.30E+07	2.00
	Combo1	alum	Staph19	5.30E+07	2.02
50	Sta011	alum	Mu-50	4.30E+07	1.33
	EsxAB	alum	Mu-50	4.30E+07	0.36
	HlaH35L	alum	Mu-50	4.30E+07	0.11
	Sta006	alum	Mu-50	4.30E+07	1.05
	Combo1	alum	Mu-50	4.30E+07	1.34
55	Sta011	alum	Staph19	4.40E+07	1.07
	EsxAB	alum	Staph19	4.40E+07	0.94

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HlaH35L	alum	Staph19	4.40E+07	1.19
Sta006	alum	Staph19	4.40E+07	2.31
Combo1	alum	Staph19	4.40E+07	2.45

* alum = aluminium hydroxide

** Log reduction in kidney CFU

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EMBODIMENTS OF THE INVENTION

35

[0650]

1. An immunogenic composition comprising a combination of antigens, said combination comprising two or more antigens selected from the group consisting of: (1) a sta006 antigen; (2) a sta011 antigen; (3) a esxA antigen; (4)
 40 a esxB antigen; (5) a hla antigen; (6) a ebpS antigen; (7) a efb antigen; (8) a emp antigen; (9) a esaC antigen; (10) a coA antigen; (11) a eap antigen; (12) a FnBA antigen; (13) a FnBB antigen; (14) a ebhA antigen; (15) a hlgB antigen; (16) a hlgC antigen; (17) a isdA antigen; (18) a isdB antigen; (19) a isdC antigen; (20) a isdG antigen; (21) a isdH antigen; (22) a isdI antigen; (23) a lukD antigen; (24) a lukE antigen; (25) a lukF antigen; (26) a lukS antigen; (27) a nuc antigen; (28) a sasA antigen; (29) a sasB antigen; (30) a sasC antigen; (31) a sasD antigen; (32) a sasF antigen; (33) a sdrC antigen; (34) a sdrD antigen; (35) a sdrE2 antigen; (36) a spa antigen; (37) a clfA antigen; (38) a clfB antigen; (39) a sta001 antigen; (40) a sta002 antigen; (41) a sta003 antigen; (42) a sta004 antigen; (43) a sta005 antigen; (44) a sta007 antigen; (45) a sta008 antigen; (46) a sta009 antigen; (47) a sta010 antigen; (48) a sta012 antigen; (49) a sta013 antigen; (50) a sta014 antigen; (51) a sta015 antigen; (52) a sta016 antigen; (53) a sta017 antigen; (54) a sta018 antigen; (55) a sta019 antigen; (56) a sta020 antigen; (57) a sta021 antigen; (58) a sta022 antigen; (59) a sta023 antigen; (60) a sta024 antigen; (61) a sta025 antigen; (62) a sta026 antigen; (63) a sta027 antigen; (64) a sta028 antigen; (65) a sta029 antigen; (66) a sta030 antigen; (67) a sta031 antigen; (68) a sta032 antigen; (69) a sta033 antigen; (70) a sta034 antigen; (71) a sta035 antigen; (72) a sta036 antigen; (73) a sta037 antigen; (74) a sta038 antigen; (75) a sta039 antigen; (76) a sta040 antigen; (77) a sta041 antigen; (78) a sta042 antigen; (79) a sta043 antigen; (80) a sta044 antigen; (81) a sta045 antigen; (82) a sta046 antigen; (83) a sta047 antigen; (84) a sta048 antigen; (85) a sta049 antigen; (86) a sta050 antigen; (87) a sta051 antigen; (88) a sta052 antigen; (89) a sta053 antigen; (90) a sta054 antigen; (91) a sta055 antigen; (92) a sta056 antigen; (93) a sta057 antigen; (94) a sta058 antigen; (95) a sta059 antigen; (96) a sta060 antigen; (97) a sta061 antigen; (98) a sta062 antigen; (99) a sta063 antigen; (100) a sta064 antigen; (101) a sta065 antigen; (102) a sta066 antigen; (103)

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a sta067 antigen; (104) a sta068 antigen; (105) a sta069 antigen; (106) a sta070 antigen; (107) a sta071 antigen; (108) a sta072 antigen; (109) a sta073 antigen; (110) a sta074 antigen; (111) a sta075 antigen; (112) a sta076 antigen; (113) a sta077 antigen; (114) a sta078 antigen; (115) a sta079 antigen; (116) a sta080 antigen; (117) a sta082 antigen; (118) a sta083 antigen; (119) a sta084 antigen; (120) a sta085 antigen; (121) a sta086 antigen; (122) a sta087 antigen; (123) a sta088 antigen; (124) a sta089 antigen; (125) a sta090 antigen; (126) a sta091 antigen; (127) a sta092 antigen; (128) a sta093 antigen; (129) a sta094 antigen; (130) a sta095 antigen; (131) a sta096 antigen; (132) a sta097 antigen; (133) a sta098 antigen; (134) a sta099 antigen; (135) a sta100 antigen; (136) a sta101 antigen; (137) a sta102 antigen; (138) a sta103 antigen; (139) a sta104 antigen; (140) a sta105 antigen; (141) a sta106 antigen; (142) a sta107 antigen; (143) a sta108 antigen; (144) a sta109 antigen; (145) a sta110 antigen; (146) a sta111 antigen; (147) a sta112 antigen; (148) a sta113 antigen; (149) a sta114 antigen; (150) a sta115 antigen; (151) a sta116 antigen; (152) a sta117 antigen; (153) a sta118 antigen; (154) a sta119 antigen; (155) a sta120 antigen; (156) a NW_6 antigen; (157) a NW_9 antigen; (158) a NW_10 antigen; (159) a NW_7 antigen; (160) a NW_8 antigen; (161) a NW_2 antigen; (162) a NW_1 antigen; (163) a sta081 antigen; and (164) a NW_5 antigen.

2. The composition of embodiment 1, comprising at least one antigen selected from numbers (3) to (38) and at least one antigen selected from numbers (1), (2) and (37) to (149).

3. The composition of embodiment 2, comprising:

at least one antigen selected from numbers (37), (38), (8), (9), (3), (4), (5), (17), (18), (19), (31), (32), (33), (34), (35) and (36);
and at least one antigen selected from (40), (1), (43), (2), (64), (96), (133) and (147).

4. The composition of embodiment 1, comprising two or more antigens selected from the group consisting of: (1) a *clfA* antigen; (2) a *clfB* antigen; (3) a *sdrE2* antigen; (4) a *sdrC* antigen; (5) a *SasF* antigen; (6) a *emp* antigen; (7) a *sdrD* antigen; (8) a *spa* antigen; (9) a *esaC* antigen; (10) a *esxA* antigen; (11) a *esxB* antigen; (12) a *sta006* antigen; (13) a *isdC* antigen; (14) a *hla* antigen; (15) a *sta011* antigen; (16) *isdA* antigen; (17) a *isdB* antigen; (18) a *sasF* antigen.

5. The composition of embodiment 1, two or more antigens selected from the group consisting of: (1) a *esxA* antigen; (2) a *esxB* antigen; (3) a *sta006* antigen; (4) a *hla* antigen; (5) a *sta011* antigen.

6. The composition of any preceding embodiment, wherein one or more of said antigens is adsorbed to an aluminium hydroxide adjuvant, and optionally wherein the composition includes a histidine buffer.

7. The composition of any preceding embodiment, further comprising: one or more conjugates of (i) a *S. aureus* exopolysaccharide and (ii) a carrier protein.

8. The composition of any preceding embodiment, further comprising: one or more conjugates of (i) a *S. aureus* capsular polysaccharide and (ii) a carrier protein.

9. A polypeptide of formula $\text{NH}_2\text{-A-}\{-\text{X-L-}\}_n\text{-B-COOH}$, wherein:

X is an amino acid sequence of a staphylococcal antigen, selected from the group consisting of *S. aureus* antigens *sta006*, *sta011*, *esxA*, *esxB*, *hla*, *clfA*, *clfB*, *coa*, *eap*, *ebhA*, *ebpS*, *efb*, *emp*, *esaC*, *FnBA*, *FnBB*, *hlgB*, *hlgC*, *isdA*, *isdB*, *isdC*, *isdG*, *isdH*, *isdI*, *lukD*, *lukE*, *lukF*, *lukS*, *nuc*, *sasA*, *sasB*, *sasC*, *sasD*, *sasF*, *sdrC*, *sdrD*, *sdrE2*, *spa*, *sta001*, *sta002*, *sta003*, *sta004*, *sta005*, *sta007*, *sta008*, *sta009*, *sta010*, *sta012*, *sta013*, *sta014*, *sta015*, *sta016*, *sta017*, *sta018*, *sta019*, *sta020*, *sta021*, *sta022*, *sta023*, *sta024*, *sta025*, *sta026*, *sta027*, *sta028*, *sta029*, *sta030*, *sta031*, *sta032*, *sta033*, *sta034*, *sta035*, *sta036*, *sta037*, *sta038*, *sta039*, *sta040*, *sta041*, *sta042*, *sta043*, *sta044*, *sta045*, *sta046*, *sta047*, *sta048*, *sta049*, *sta050*, *sta051*, *sta052*, *sta053*, *sta054*, *sta055*, *sta056*, *sta057*, *sta058*, *sta059*, *sta060*, *sta061*, *sta062*, *sta063*, *sta064*, *sta065*, *sta066*, *sta067*, *sta068*, *sta069*, *sta070*, *sta071*, *sta072*, *sta073*, *sta074*, *sta075*, *sta076*, *sta077*, *sta078*, *sta079*, *sta080*, *sta081*, *sta082*, *sta083*, *sta084*, *sta085*, *sta086*, *sta087*, *sta088*, *sta089*, *sta090*, *sta091*, *sta092*, *sta093*, *sta094*, *sta095*, *sta096*, *sta097*, *sta098*, *sta099*, *sta100*, *sta101*, *sta102*, *sta103*, *sta104*, *sta105*, *sta106*, *sta107*, *sta108*, *sta109*, *sta110*, *sta111*, *sta112*, *sta113*, *sta114*, *sta115*, *sta116*, *sta117*, *sta118*, *NW_6*, *NW_9*, *NW_10*, *NW_7*, *NW_8*, *NW_2*, *NW_1*, and *NW_5*;

L is an optional linker amino acid sequence;

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A is an optional N-terminal amino acid sequence;
B is an optional C-terminal amino acid sequence; and
n is an integer of 2 or more.

5 10. An immunogenic composition comprising the polypeptide of embodiment 9 and further comprising: (A) one or more conjugates of (i) a *S.aureus* exopolysaccharide and (ii) a carrier protein; and/or (B) one or more conjugates of (i) a *S.aureus* capsular polysaccharide and (ii) a carrier protein.

10 11. The composition or polypeptide of any preceding embodiment, wherein the clfA antigen can elicit an antibody which recognises SEQ ID NO: 1 and comprises an amino acid sequence: (a) having 80% or more identity to SEQ ID NO: 1; and/or (b) comprising a fragment of at least 7 consecutive amino acids of SEQ ID NO: 1, wherein the fragment comprises an epitope from SEQ ID NO: 1.

15 12. A polypeptide comprising amino acid sequence (a) having 80% or more identity to SEQ ID NO: 151; and/or (b) comprising a fragment of at least 7 consecutive amino acids from amino acids 1-97 of SEQ ID NO: 151 and at least 7 consecutive amino acids from amino acids 104-207 of SEQ ID NO: 151, wherein the polypeptide can elicit antibodies which recognise both the wild-type staphylococcal protein comprising SEQ ID NO: 10 and the wild-type staphylococcal protein comprising SEQ ID NO: 11.

20 13. An immunogenic composition comprising the polypeptide of embodiment 12 and one or more of (i) a sta006 antigen; (ii) a hla antigen; and/or (iii) a sta011 antigen.

14. The composition of embodiment 13, including an adjuvant.

25 15. A polypeptide comprising amino acid sequence having 80% or more identity to an amino acid sequence selected from SEQ ID NOs: 151, 152, 168, 202, 203, 204, 205, 206, 207, 208, 209, 210,211,212,220,221,222,223,224,237,238,241.

30 16. A polypeptide comprising an amino acid sequence selected from SEQ ID NOs: 151, 152, 168, 202, 203, 204, 205, 206, 207, 208, 209, 210, 211, 212, 220, 221, 222, 223, 224, 237, 238, 241, 242,243,244,245.

35 17. A polypeptide comprising: (a) a first sequence having 90% or more identity to SEQ ID NO: 218; and (b) a second sequence having 90% or more identity to SEQ ID NO: 219, wherein the first and second sequences are either directly joined or are joined by an intervening amino acid sequence having fewer than 10 amino acids.

18. A pharmaceutical composition comprising the polypeptide of any one of embodiments 12, 15, 16 or 17.

40 19. A method for raising an immune response in a mammal comprising the step of administering to the mammal an effective amount of the polypeptide or composition of any preceding embodiment.

45 20. Nucleic acid encoding the polypeptide of embodiment 9, 12, 15, 16 or 17.

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SEQUENCE LISTING

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<130> P054462W0

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 Ile Tyr Ala Ile Gly Leu Thr Asn Lys Tyr Glu Tyr Gly Asp Asn Ile
 65 70 75 80
 Tyr Lys Glu Ala Lys Asp Arg Leu Leu Glu Lys Val Leu Arg Glu Asp
 85 90 95
 Gln Tyr Leu Leu Glu Arg Lys Lys Ser Gln Tyr Glu Asp Tyr Lys Gln
 100 105 110
 Trp Tyr Ala Asn Tyr Lys Lys Glu Asn Pro Arg Thr Asp Leu Lys Met
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 Ala Asn Phe His Lys Tyr Asn Leu Glu Glu Leu Ser Met Lys Glu Tyr
 130 135 140
 Asn Glu Leu Gln Asp Ala Leu Lys Arg Ala Leu Asp Asp Phe His Arg
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 Glu Val Lys Asp Ile Lys Asp Lys Asn Ser Asp Leu Lys Thr Phe Asn
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 Ala Ala Glu Glu Asp Lys Ala Thr Lys Glu Val Tyr Asp Leu Val Ser
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 Glu Ile Asp Thr Leu Val Val Ser Tyr Tyr Gly Asp Lys Asp Tyr Gly
 195 200 205
 Glu His Ala Lys Glu Leu Arg Ala Lys Leu Asp Leu Ile Leu Gly Asp
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Thr Asp Asn Pro His Lys Ile Thr Asn Glu Arg Ile Lys Lys Glu Met
 225 230 235 240
 5 Ile Asp Asp Leu Asn Ser Ile Ile Asp Asp Phe Phe Met Glu Thr Lys
 245 250 255
 Gln Asn Arg Pro Lys Ser Ile Thr Lys Tyr Asn Pro Thr Thr His Asn
 260 265 270
 10 Tyr Lys Thr Asn Ser Asp Asn Lys Pro Asn Phe Asp Lys Leu Val Glu
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 Glu Thr Lys Lys Ala Val Lys Glu Ala Asp Asp Ser Trp Lys Lys Lys
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 15 Thr Val Lys Lys Tyr Gly Glu Thr Glu Thr Lys Ser Pro Val Val Lys
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 Glu Glu Lys Lys Val Glu Glu Pro Gln Ala Pro Lys Val Asp Asn Gln
 325 330 335
 20 Gln Glu Val Lys Thr Thr Ala Gly Lys Ala Glu Glu Thr Thr Gln Pro
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 Val Ala Gln Pro Leu Val Lys Ile Pro Gln Gly Thr Ile Thr Gly Glu
 355 360 365
 25 Ile Val Lys Gly Pro Glu Tyr Pro Thr Met Glu Asn Lys Thr Val Gln
 370 375 380
 Gly Glu Ile Val Gln Gly Pro Asp Phe Leu Thr Met Glu Gln Ser Gly
 385 390 395 400
 Pro Ser Leu Ser Asn Asn Tyr Thr Asn Pro Pro Leu Thr Asn Pro Ile
 405 410 415
 30 Leu Glu Gly Leu Glu Gly Ser Ser Ser Lys Leu Glu Ile Lys Pro Gln
 420 425 430
 Gly Thr Glu Ser Thr Leu Lys Gly Thr Gln Gly Glu Ser Ser Asp Ile
 435 440 445
 35 Glu Val Lys Pro Gln Ala Thr Glu Thr Thr Glu Ala Ser Gln Tyr Gly
 450 455 460
 Pro Arg Pro Gln Phe Asn Lys Thr Pro Lys Tyr Val Lys Tyr Arg Asp
 465 470 475 480
 40 Ala Gly Thr Gly Ile Arg Glu Tyr Asn Asp Gly Thr Phe Gly Tyr Glu
 485 490 495
 Ala Arg Pro Arg Phe Asn Lys Pro Ser Glu Thr Asn Ala Tyr Asn Val
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 45 Thr Thr His Ala Asn Gly Gln Val Ser Tyr Gly Ala Arg Pro Thr Tyr
 515 520 525
 Lys Lys Pro Ser Glu Thr Asn Ala Tyr Asn Val Thr Thr His Ala Asn
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 50 Gly Gln Val Ser Tyr Gly Ala Arg Pro Thr Gln Asn Lys Pro Ser Lys
 545 550 555 560
 Thr Asn Ala Tyr Asn Val Thr Thr His Gly Asn Gly Gln Val Ser Tyr
 565 570 575
 55 Gly Ala Arg Pro Thr Gln Asn Lys Pro Ser Lys Thr Asn Ala Tyr Asn
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 Val Thr Thr His Ala Asn Gly Gln Val Ser Tyr Gly Ala Arg Pro Thr

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	1				5					10					15		
15	Ala	Ser	Thr	Gly	Ala	Asn	Phe	Asn	Asn	Asn	Glu	Ala	Ser	Ala	Ala	Ala	
				20					25					30			
	Lys	Pro	Leu	Asp	Lys	Ser	Ser	Ser	Ser	Leu	His	His	Gly	Tyr	Ser	Lys	
			35					40					45				
20	Val	His	Val	Pro	Tyr	Ala	Ile	Thr	Val	Asn	Gly	Thr	Ser	Gln	Asn	Ile	
		50					55					60					
	Leu	Ser	Ser	Leu	Thr	Phe	Asn	Lys	Asn	Gln	Asn	Ile	Ser	Tyr	Lys	Asp	
	65					70					75					80	
25	Leu	Glu	Asp	Arg	Val	Lys	Ser	Val	Leu	Lys	Ser	Asp	Arg	Gly	Ile	Ser	
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	Asp	Ile	Asp	Leu	Arg	Leu	Ser	Lys	Gln	Ala	Lys	Tyr	Thr	Val	Tyr	Phe	
				100					105					110			
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	Gln	Val	Pro	Tyr	Thr	Ile	Thr	Val	Asn	Gly	Thr	Ser	Gln	Asn	Ile	Leu	
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40	Ser	Asn	Leu	Thr	Phe	Asn	Lys	Asn	Gln	Asn	Ile	Ser	Tyr	Lys	Asp	Leu	
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	Tyr	Ser	Ile	Asn	Leu	Asn	Gly	Thr	Ser	Thr	Asn	Ile	Leu	Ser	Asn	Leu	
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Ile Lys Ser Val Leu Lys His Asp Arg Gly Ile Ser Glu Gln Asp Leu
 305 310 315 320

Lys Tyr Ala Lys Lys Ala Tyr Tyr Thr Val Tyr Phe Lys Asn Gly Gly
 5 325 330 335

Lys Arg Ile Leu Gln Leu Asn Ser Lys Asn Tyr Thr Ala Asn Leu Val
 340 345 350

His Ala Lys Asp Val Lys Arg Ile Glu Ile Thr Val Lys Thr Gly Thr
 10 355 360 365

Lys Ala Lys Ala Asp Arg Tyr Val Pro Tyr Thr Ile Ala Val Asn Gly
 370 375 380

Thr Ser Thr Pro Ile Leu Ser Asp Leu Lys Phe Thr Gly Asp Pro Arg
 15 385 390 395 400

Val Gly Tyr Lys Asp Ile Ser Lys Lys Val Lys Ser Val Leu Lys His
 405 410 415

Asp Arg Gly Ile Gly Glu Arg Glu Leu Lys Tyr Ala Lys Lys Ala Thr
 20 420 425 430

Tyr Thr Val His Phe Lys Asn Gly Thr Lys Lys Val Ile Asn Ile Asn
 435 440 445

Ser Asn Ile Ser Gln Leu Asn Leu Leu Tyr Val Gln Asp Ile Lys Lys
 25 450 455 460

Ile Asp Ile Asp Val Lys Thr Gly Thr Lys Ala Lys Ala Asp Ser Tyr
 465 470 475 480

Val Pro Tyr Thr Ile Ala Val Asn Gly Thr Ser Thr Pro Ile Leu Ser
 30 485 490 495

Lys Leu Lys Ile Ser Asn Lys Gln Leu Ile Ser Tyr Lys Tyr Leu Asn
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Asp Lys Val Lys Ser Val Leu Lys Ser Glu Arg Gly Ile Ser Asp Leu
 35 515 520 525

Asp Leu Lys Phe Ala Lys Gln Ala Lys Tyr Thr Val Tyr Phe Lys Asn
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Gly Lys Lys Gln Val Val Asn Leu Lys Ser Asp Ile Phe Thr Pro Asn
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Tyr Thr Lys Ser Lys Lys Asn Lys
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55 Asn Thr Ser Gln Ala His Ala Ala Glu Thr Asn Gln Pro Ala Ser Val
 35 40 45

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Val Lys Gln Lys Gln Gln Ser Asn Asn Glu Gln Thr Glu Asn Arg Glu
 50 55 60
 Ser Gln Val Gln Asn Ser Gln Asn Ser Gln Asn Gly Gln Ser Leu Ser
 65 70 75 80
 Ala Thr His Glu Asn Glu Gln Pro Asn Ile Ser Gln Ala Asn Leu Val
 85 90 95
 Asp Gln Lys Val Ala Gln Ser Ser Thr Thr Asn Asp Glu Gln Pro Ala
 100 105 110
 Ser Gln Asn Val Asn Thr Lys Lys Asp Ser Ala Thr Ala Ala Thr Thr
 115 120 125
 Gln Pro Asp Lys Glu Gln Ser Lys His Lys Gln Asn Glu Ser Gln Ser
 130 135 140
 Ala Asn Lys Asn Gly Asn Asp Asn Arg Ala Ala His Val Glu Asn His
 145 150 155 160
 Glu Ala Asn Val Val Thr Ala Ser Asp Ser Ser Asp Asn Gly Asn Val
 165 170 175
 Gln His Asp Arg Asn Glu Leu Gln Ala Phe Phe Asp Ala Asn Tyr His
 180 185 190
 Asp Tyr Arg Phe Ile Asp Arg Glu Asn Ala Asp Ser Gly Thr Phe Asn
 195 200 205
 Tyr Val Lys Gly Ile Phe Asp Lys Ile Asn Thr Leu Leu Gly Ser Asn
 210 215 220
 Asp Pro Ile Asn Asn Lys Asp Leu Gln Leu Ala Tyr Lys Glu Leu Glu
 225 230 235 240
 Gln Ala Val Ala Leu Ile Arg Thr Met Pro Gln Arg Gln Gln Thr Ser
 245 250 255
 Arg Arg Ser Asn Arg Ile Gln Thr Arg Ser Val Glu Ser Arg Ala Ala
 260 265 270
 Glu Pro Arg Ser Val Ser Asp Tyr Gln Asn Ala Asn Ser Ser Tyr Tyr
 275 280 285
 Val Glu Asn Ala Asn Asp Gly Ser Gly Tyr Pro Val Gly Thr Tyr Ile
 290 295 300
 Asn Ala Ser Ser Lys Gly Ala Pro Tyr Asn Leu Pro Thr Thr Pro Trp
 305 310 315 320
 Asn Thr Leu Lys Ala Ser Asp Ser Lys Glu Ile Ala Leu Met Thr Ala
 325 330 335
 Lys Gln Thr Gly Asp Gly Tyr Gln Trp Val Ile Lys Phe Asn Lys Gly
 340 345 350
 His Ala Pro His Gln Asn Met Ile Phe Trp Phe Ala Leu Pro Ala Asp
 355 360 365
 Gln Val Pro Val Gly Arg Thr Asp Phe Val Thr Val Asn Ser Asp Gly
 370 375 380
 Thr Asn Val Gln Trp Ser His Gly Ala Gly Ala Gly Ala Asn Lys Pro
 385 390 395 400
 Leu Gln Gln Met Trp Glu Tyr Gly Val Asn Asp Pro His Arg Ser His
 405 410 415
 Asp Phe Lys Ile Arg Asn Arg Ser Gly Gln Val Ile Tyr Asp Trp Pro

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			420					425				430				
	Thr	Val	His	Ile	Tyr	Ser	Leu	Glu	Asp	Leu	Ser	Arg	Ala	Ser	Asp	Tyr
			435					440					445			
5	Phe	Ser	Glu	Ala	Gly	Ala	Thr	Pro	Ala	Thr	Lys	Ala	Phe	Gly	Arg	Gln
		450					455					460				
	Asn	Phe	Glu	Tyr	Ile	Asn	Gly	Gln	Lys	Pro	Ala	Glu	Ser	Pro	Gly	Val
	465					470					475					480
10	Pro	Lys	Val	Tyr	Thr	Phe	Ile	Gly	Gln	Gly	Asp	Ala	Ser	Tyr	Thr	Ile
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	Ser	Phe	Lys	Thr	Gln	Gly	Pro	Thr	Val	Asn	Lys	Leu	Tyr	Tyr	Ala	Ala
				500					505					510		
15	Gly	Gly	Arg	Ala	Leu	Glu	Tyr	Asn	Gln	Leu	Phe	Met	Tyr	Ser	Gln	Leu
			515					520					525			
	Tyr	Val	Glu	Ser	Thr	Gln	Asp	His	Gln	Gln	Arg	Leu	Asn	Gly	Leu	Arg
		530					535					540				
20	Gln	Val	Val	Asn	Arg	Thr	Tyr	Arg	Ile	Gly	Thr	Thr	Lys	Arg	Val	Glu
	545					550					555					560
	Val	Ser	Gln	Gly	Asn	Val	Gln	Thr	Lys	Lys	Val	Leu	Glu	Ser	Thr	Asn
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25	Leu	Asn	Ile	Asp	Asp	Phe	Val	Asp	Asp	Pro	Leu	Ser	Tyr	Val	Lys	Thr
				580				585						590		
	Pro	Ser	Asn	Lys	Val	Leu	Gly	Phe	Tyr	Ser	Asn	Asn	Ala	Asn	Thr	Asn
			595					600					605			
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	Gln	Leu	Phe	Thr	Asp	Gln	Lys	Leu	Gln	Glu	Ala	Ala	Arg	Thr	Arg	Asn
	625					630					635					640
35	Pro	Ile	Arg	Leu	Met	Ile	Gly	Phe	Asp	Tyr	Pro	Asp	Ala	Tyr	Gly	Asn
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	Ser	Glu	Thr	Leu	Val	Pro	Val	Asn	Leu	Thr	Val	Leu	Pro	Glu	Ile	Gln
				660					665					670		
40	His	Asn	Ile	Lys	Phe	Phe	Lys	Asn	Asp	Asp	Thr	Gln	Asn	Ile	Ala	Glu
			675					680					685			
	Lys	Pro	Phe	Ser	Lys	Gln	Ala	Gly	His	Pro	Val	Phe	Tyr	Val	Tyr	Ala
		690					695					700				
45	Gly	Asn	Gln	Gly	Asn	Ala	Ser	Val	Asn	Leu	Gly	Gly	Ser	Val	Thr	Ser
	705					710					715					720
	Ile	Gln	Pro	Leu	Arg	Ile	Asn	Leu	Thr	Ser	Asn	Glu	Asn	Phe	Thr	Asp
					725					730					735	
50	Lys	Asp	Trp	Gln	Ile	Thr	Gly	Ile	Pro	Arg	Thr	Leu	His	Ile	Glu	Asn
				740					745					750		
	Ser	Thr	Asn	Arg	Pro	Asn	Asn	Ala	Arg	Glu	Arg	Asn	Ile	Glu	Leu	Val
			755					760					765			
55	Gly	Asn	Leu	Leu	Pro	Gly	Asp	Tyr	Phe	Gly	Thr	Ile	Arg	Phe	Gly	Arg
		770					775					780				
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	785					790					795					800

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Thr Thr Thr Ala Glu Gln Leu Arg Gly Thr Ala Leu Gln Lys Val Pro
 805 810 815
 Val Asn Ile Ser Gly Ile Pro Leu Asp Pro Ser Ala Leu Val Tyr Leu
 5 820 825 830
 Val Ala Pro Thr Asn Gln Thr Thr Asn Gly Gly Ser Glu Ala Asp Gln
 835 840 845
 Ile Pro Ser Gly Tyr Thr Ile Leu Ala Thr Gly Thr Pro Asp Gly Val
 10 850 855 860
 His Asn Thr Ile Thr Ile Arg Pro Gln Asp Tyr Val Val Phe Ile Pro
 865 870 875 880
 Pro Val Gly Lys Gln Ile Arg Ala Val Val Tyr Tyr Asn Lys Val Val
 15 885 890 895
 Ala Ser Asn Met Ser Asn Ala Val Thr Ile Leu Pro Asp Asp Ile Pro
 900 905
 Pro Thr Ile Asn Asn Pro Val Gly Ile Asn Ala Lys Tyr Tyr Arg Gly
 20 915 920 925
 Asp Glu Val Asn Phe Thr Met Gly Val Ser Asp Arg His Ser Gly Ile
 930 935 940
 Lys Asn Thr Thr Ile Thr Thr Leu Pro Asn Gly Trp Thr Ser Asn Leu
 25 945 950 955 960
 Thr Lys Ala Asp Lys Asn Asn Gly Ser Leu Ser Ile Thr Gly Arg Val
 965 970 975
 Ser Met Asn Gln Ala Phe Asn Ser Asp Ile Thr Phe Lys Val Ser Ala
 30 980 985 990
 Thr Asp Asn Val Asn Asn Thr Thr Asn Asp Ser Gln Ser Lys His Val
 995 1000 1005
 Ser Ile His Val Gly Lys Ile Ser Glu Asp Ala His Pro Ile Val Leu
 35 1010 1015 1020
 Gly Asn Thr Glu Lys Val Val Val Val Asn Pro Thr Ala Val Ser Asn
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 Asp Glu Lys Gln Ser Ile Ile Thr Ala Phe Met Asn Lys Asn Gln Asn
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 Ile Arg Gly Tyr Leu Ala Ser Thr Asp Pro Val Thr Val Asp Asn Asn
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 Gly Asn Val Thr Leu His Tyr Arg Asp Gly Ser Ser Thr Thr Leu Asp
 45 1075 1080 1085
 Ala Thr Asn Val Met Thr Tyr Glu Pro Val Val Lys Pro Glu Tyr Gln
 1090 1095 1100
 Thr Val Asn Ala Ala Lys Thr Ala Thr Val Thr Ile Ala Lys Gly Gln
 50 1105 1110 1115 1120
 Ser Phe Ser Ile Gly Asp Ile Lys Gln Tyr Phe Thr Leu Ser Asn Gly
 1125 1130 1135
 Gln Pro Ile Pro Ser Gly Thr Phe Thr Asn Ile Thr Ser Asp Arg Thr
 1140 1145 1150
 Ile Pro Thr Ala Gln Glu Val Ser Gln Met Asn Ala Gly Thr Gln Leu
 55 1155 1160 1165

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Tyr His Ile Thr Ala Thr Asn Ala Tyr His Lys Asp Ser Glu Asp Phe
 1170 1175 1180
 Tyr Ile Ser Leu Lys Ile Ile Asp Val Lys Gln Pro Glu Gly Asp Gln
 5 1185 1190 1195
 Arg Val Tyr Arg Thr Ser Thr Tyr Asp Leu Thr Thr Asp Glu Ile Ser
 1205 1210 1215
 Lys Val Lys Gln Ala Phe Ile Asn Ala Asn Arg Asp Val Ile Thr Leu
 10 1220 1225 1230
 Ala Glu Gly Asp Ile Ser Val Thr Asn Thr Pro Asn Gly Ala Asn Val
 1235 1240 1245
 Ser Thr Ile Thr Val Asn Ile Asn Lys Gly Arg Leu Thr Lys Ser Phe
 15 1250 1255 1260
 Ala Ser Asn Leu Ala Asn Met Asn Phe Leu Arg Trp Val Asn Phe Pro
 1265 1270 1275 1280
 Gln Asp Tyr Thr Val Thr Trp Thr Asn Ala Lys Ile Ala Asn Arg Pro
 20 1285 1290 1295
 Thr Asp Gly Gly Leu Ser Trp Ser Asp Asp His Lys Ser Leu Ile Tyr
 1300 1305 1310
 Arg Tyr Asp Ala Thr Leu Gly Thr Gln Ile Thr Thr Asn Asp Ile Leu
 25 1315 1320 1325
 Thr Met Leu Lys Ala Thr Thr Thr Val Pro Gly Leu Arg Asn Asn Ile
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 Thr Gly Asn Glu Lys Ser Gln Ala Glu Ala Gly Gly Arg Pro Asn Phe
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 Arg Thr Thr Gly Tyr Ser Gln Ser Asn Ala Thr Thr Asp Gly Gln Arg
 30 1365 1370 1375
 Gln Phe Thr Leu Asn Gly Gln Val Ile Gln Val Leu Asp Ile Ile Asn
 1380 1385 1390
 Pro Ser Asn Gly Tyr Gly Gly Gln Pro Val Thr Asn Ser Asn Thr Arg
 35 1395 1400 1405
 Ala Asn His Ser Asn Ser Thr Val Val Asn Val Asn Glu Pro Ala Ala
 1410 1415 1420
 Asn Gly Ala Gly Ala Phe Thr Ile Asp His Val Val Lys Ser Asn Ser
 40 1425 1430 1435 1440
 Thr His Asn Ala Ser Asp Ala Val Tyr Lys Ala Gln Leu Tyr Leu Thr
 1445 1450 1455
 Pro Tyr Gly Pro Lys Gln Tyr Val Glu His Leu Asn Gln Asn Thr Gly
 45 1460 1465 1470
 Asn Thr Thr Asp Ala Ile Asn Ile Tyr Phe Val Pro Ser Asp Leu Val
 1475 1480 1485
 Asn Pro Thr Ile Ser Val Gly Asn Tyr Thr Asn His Gln Val Phe Ser
 50 1490 1495 1500
 Gly Glu Thr Phe Thr Asn Thr Ile Thr Ala Asn Asp Asn Phe Gly Val
 1505 1510 1515 1520
 Gln Ser Val Thr Val Pro Asn Thr Ser Gln Ile Thr Gly Thr Val Asp
 55 1525 1530 1535
 Asn Asn His Gln His Val Ser Ala Thr Ala Pro Asn Val Thr Ser Ala

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	Thr Asn Lys Thr Ile Asn Leu Leu Ala Thr Asp Thr Ser Gly Asn Thr 1555 1560 1565		
5	Ala Thr Thr Ser Phe Asn Val Thr Val Lys Pro Leu Arg Asp Lys Tyr 1570 1575 1580		
	Arg Val Gly Thr Ser Ser Thr Ala Ala Asn Pro Val Arg Ile Ala Asn 1585 1590 1595 1600		
10	Ile Ser Asn Asn Ala Thr Val Ser Gln Ala Asp Gln Thr Thr Ile Ile 1605 1610 1615		
	Asn Ser Leu Thr Phe Thr Glu Thr Val Pro Asn Arg Ser Tyr Ala Arg 1620 1625 1630		
15	Ala Ser Ala Asn Glu Ile Thr Ser Lys Thr Val Ser Asn Val Ser Arg 1635 1640 1645		
	Thr Gly Asn Asn Ala Asn Val Thr Val Thr Val Thr Tyr Gln Asp Gly 1650 1655 1660		
20	Thr Thr Ser Thr Val Thr Val Pro Val Lys His Val Ile Pro Glu Ile 1665 1670 1675 1680		
	Val Ala His Ser His Tyr Thr Val Gln Gly Gln Asp Phe Pro Ala Gly 1685 1690 1695		
25	Asn Gly Ser Ser Ala Ser Asp Tyr Phe Lys Leu Ser Asn Gly Ser Asp 1700 1705 1710		
	Ile Ala Asp Ala Thr Ile Thr Trp Val Ser Gly Gln Ala Pro Asn Lys 1715 1720 1725		
30	Asp Asn Thr Arg Ile Gly Glu Asp Ile Thr Val Thr Ala His Ile Leu 1730 1735 1740		
	Ile Asp Gly Glu Thr Thr Pro Ile Thr Lys Thr Ala Thr Tyr Lys Val 1745 1750 1755 1760		
35	Val Arg Thr Val Pro Lys His Val Phe Glu Thr Ala Arg Gly Val Leu 1765 1770 1775		
	Tyr Pro Gly Val Ser Asp Met Tyr Asp Ala Lys Gln Tyr Val Lys Pro 1780 1785 1790		
40	Val Asn Asn Ser Trp Ser Thr Asn Ala Gln His Met Asn Phe Gln Phe 1795 1800 1805		
	Val Gly Thr Tyr Gly Pro Asn Lys Asp Val Val Gly Ile Ser Thr Arg 1810 1815 1820		
45	Leu Ile Arg Val Thr Tyr Asp Asn Arg Gln Thr Glu Asp Leu Thr Ile 1825 1830 1835 1840		
	Leu Ser Lys Val Lys Pro Asp Pro Pro Arg Ile Asp Ala Asn Ser Val 1845 1850 1855		
50	Thr Tyr Lys Ala Gly Leu Thr Asn Gln Glu Ile Lys Val Asn Asn Val 1860 1865 1870		
	Leu Asn Asn Ser Ser Val Lys Leu Phe Lys Ala Asp Asn Thr Pro Leu 1875 1880 1885		
55	Asn Val Thr Asn Ile Thr His Gly Ser Gly Phe Ser Ser Val Val Thr 1890 1895 1900		
	Val Ser Asp Ala Leu Pro Asn Gly Gly Ile Lys Ala Lys Ser Ser Ile 1905 1910 1915 1920		

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Ser Met Asn Asn Val Thr Tyr Thr Thr Gln Asp Glu His Gly Gln Val
 1925 1930 1935
 Val Thr Val Thr Arg Asn Glu Ser Val Asp Ser Asn Asp Ser Ala Thr
 5 1940 1945 1950
 Val Thr Val Thr Pro Gln Leu Gln Ala Thr Thr Glu Gly Ala Val Phe
 1955 1960 1965
 Ile Lys Gly Gly Asp Gly Phe Asp Phe Gly His Val Glu Arg Phe Ile
 10 1970 1975 1980
 Gln Asn Pro Pro His Gly Ala Thr Val Ala Trp His Asp Ser Pro Asp
 1985 1990 1995 2000
 Thr Trp Lys Asn Thr Val Gly Asn Thr His Lys Thr Ala Val Val Thr
 15 2005 2010 2015
 Leu Pro Asn Gly Gln Gly Thr Arg Asn Val Glu Val Pro Val Lys Val
 2020 2025 2030
 Tyr Pro Val Ala Asn Ala Lys Ala Pro Ser Arg Asp Val Lys Gly Gln
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 Asn Leu Thr Asn Gly Thr Asp Ala Met Asn Tyr Ile Thr Phe Asp Pro
 2050 2055 2060
 Asn Thr Asn Thr Asn Gly Ile Thr Ala Ala Trp Ala Asn Arg Gln Gln
 25 2065 2070 2075 2080
 Pro Asn Asn Gln Gln Ala Gly Val Gln His Leu Asn Val Asp Val Thr
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 Tyr Pro Gly Ile Ser Ala Ala Lys Arg Val Pro Val Thr Val Asn Val
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 Tyr Gln Phe Glu Phe Pro Gln Thr Thr Tyr Thr Thr Thr Val Gly Gly
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 Thr Leu Ala Ser Gly Thr Gln Ala Ser Gly Tyr Ala His Met Gln Asn
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 Ala Thr Gly Leu Pro Thr Asp Gly Phe Thr Tyr Lys Trp Asn Arg Asp
 2145 2150 2155 2160
 Thr Thr Gly Thr Asn Asp Ala Asn Trp Ser Ala Met Asn Lys Pro Asn
 40 2165 2170 2175
 Val Ala Lys Val Val Asn Ala Lys Tyr Asp Val Ile Tyr Asn Gly His
 2180 2185 2190
 Thr Phe Ala Thr Ser Leu Pro Ala Lys Phe Val Val Lys Asp Val Gln
 45 2195 2200 2205
 Pro Ala Lys Pro Thr Val Thr Glu Thr Ala Ala Gly Ala Ile Thr Ile
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 Thr Tyr Ala Asp Lys Leu Val Ile Lys Arg Asn Gly Asn Val Val Thr
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 Thr Phe Thr Arg Arg Asn Asn Thr Ser Pro Trp Val Lys Glu Ala Ser
 55 2260 2265 2270
 Ala Ala Thr Val Ala Gly Ile Ala Gly Thr Asn Asn Gly Ile Thr Val
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Ala Ala Gly Thr Phe Asn Pro Ala Asp Thr Ile Gln Val Val Ala Thr
 2290 2295 2300

5 Gln Gly Ser Gly Glu Thr Val Ser Asp Glu Gln Arg Ser Asp Asp Phe
 2305 2310 2315 2320

Thr Val Val Ala Pro Gln Pro Asn Gln Ala Thr Thr Lys Ile Trp Gln
 2325 2330 2335

10 Asn Gly His Ile Asp Ile Thr Pro Asn Asn Pro Ser Gly His Leu Ile
 2340 2345 2350

Asn Pro Thr Gln Ala Met Asp Ile Ala Tyr Thr Glu Lys Val Gly Asn
 2355 2360 2365

15 Gly Ala Glu His Ser Lys Thr Ile Asn Val Val Arg Gly Gln Asn Asn
 2370 2375 2380

Gln Trp Thr Ile Ala Asn Lys Pro Asp Tyr Val Thr Leu Asp Ala Gln
 2385 2390 2395

20 Thr Gly Lys Val Thr Phe Asn Ala Asn Thr Ile Lys Pro Asn Ser Ser
 2405 2410 2415

Ile Thr Ile Thr Pro Lys Ala Gly Thr Gly His Ser Val Ser Ser Asn
 2420 2425 2430

25 Pro Ser Thr Leu Thr Ala Pro Ala Ala His Thr Val Asn Thr Thr Glu
 2435 2440 2445

Ile Val Lys Asp Tyr Gly Ser Asn Val Thr Ala Ala Glu Ile Asn Asn
 2450 2455 2460

30 Ala Val Gln Val Ala Asn Lys Arg Thr Ala Thr Ile Lys Asn Gly Thr
 2465 2470 2475 2480

Ala Met Pro Thr Asn Leu Ala Gly Gly Ser Thr Thr Thr Ile Pro Val
 2485 2490 2495

Thr Val Thr Tyr Asn Asp Gly Ser Thr Glu Glu Val Gln Glu Ser Ile
 2500 2505 2510

35 Phe Thr Lys Ala Asp Lys Arg Glu Leu Ile Thr Ala Lys Asn His Leu
 2515 2520 2525

Asp Asp Pro Val Ser Thr Glu Gly Lys Lys Pro Gly Thr Ile Thr Gln
 2530 2535 2540

40 Tyr Asn Asn Ala Met His Asn Ala Gln Gln Gln Ile Asn Thr Ala Lys
 2545 2550 2555

Thr Glu Ala Gln Gln Val Ile Asn Asn Glu Arg Ala Thr Pro Gln Gln
 2565 2570 2575

45 Val Ser Asp Ala Leu Thr Lys Val Arg Ala Ala Gln Thr Lys Ile Asp
 2580 2585 2590

Gln Ala Lys Ala Leu Leu Gln Asn Lys Glu Asp Asn Ser Gln Leu Val
 2595 2600 2605

50 Thr Ser Lys Asn Asn Leu Gln Ser Ser Val Asn Gln Val Pro Ser Thr
 2610 2615 2620

Ala Gly Met Thr Gln Gln Ser Ile Asp Asn Tyr Asn Ala Lys Lys Arg
 2625 2630 2635 2640

55 Glu Ala Glu Thr Glu Ile Thr Ala Ala Gln Arg Val Ile Asp Asn Gly
 2645 2650 2655

Asp Ala Thr Ala Gln Gln Ile Ser Asp Glu Lys His Arg Val Asp Asn

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	2660	2665	2670
	Ala Leu Thr 2675	Ala Lys His 2680	Leu Thr Ala Asp Thr 2685
5	His Ala Leu 2690	Gln Ala Val 2695	Gln Leu Asn Arg Thr 2700
	Thr Gly Lys 2705	Lys Pro Ala Ser 2710	Ile Thr Ala Tyr Asn 2715
10	Ala Leu Gln 2725	Ser Asp Leu Thr 2725	Ser Ala Lys Asn Ser 2730
	Ile Gln Lys 2740	Pro Ile Arg Thr 2740	Val Gln Glu Val Gln 2745
15	Asn Val Asn 2755	Arg Val Asn Glu 2760	Arg Leu Thr Gln Ala 2765
	Val Pro Leu 2770	Ala Asp Asn Ser 2775	Ala Leu Lys Thr Ala 2780
20	Asp Glu Glu 2785	Ile Asn Lys Ser 2790	Val Thr Thr Asp Gly 2795
	Ser Ile Gln 2805	Ala Tyr Glu Asn 2805	Ala Lys Arg Ala Gly 2810
25	Thr Asn Ala 2820	Gln Asn Val Ile 2820	Asn Asn Gly Asp Ala 2825
	Ile Ala Ala 2835	Glu Lys Thr Lys 2840	Val Glu Glu Lys Tyr 2845
30	Gln Ala Ile 2850	Ala Gly Leu Thr 2855	Pro Asp Leu Ala Pro 2860
	Lys Thr Gln 2865	Leu Gln Asn Asp 2870	Ile Asp Gln Pro Thr 2875
35	Met Thr Ser 2885	Ala Ser Ile Ala 2885	Ala Phe Asn Glu Lys 2890
	Arg Thr Lys 2900	Ile Gln Glu Ile 2900	Asp Arg Val Leu Ala 2905
40	Val Ala Thr 2915	Ile Arg Gln Asn 2920	Val Thr Ala Ala Asn 2925
	Ala Leu Asp 2930	Gln Ala Arg Asn 2935	Gly Leu Thr Val Asp 2940
45	Glu Asn Ala 2945	Lys Asn Gln Leu 2950	Gln His Ser Ile Asp 2955
	Thr Thr Gly 2965	Met Thr Gln Asp 2965	Ser Ile Asn Ala Tyr 2970
50	Thr Ala Ala 2980	Arg Asn Lys Ile 2980	Gln Gln Ile Asn Gln 2985
	Ser Pro Thr 2995	Val Glu Gln Ile 3000	Asn Thr Asn Thr Ser 3005
55	Ala Lys Ser 3010	Asp Leu Asp His 3015	Ala Arg Gln Ala Leu 3020
	Ala Pro Leu 3025	Gln Thr Ala Lys 3030	Thr Gln Leu Glu Gln 3035
			Ser Ile Asn Gln 3040

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Pro Thr Asp Thr Thr Gly Met Thr Thr Ala Ser Leu Asn Ala Tyr Asn
 3045 3050 3055
 5 Gln Lys Leu Gln Ala Ala Arg Gln Lys Leu Thr Glu Ile Asn Gln Val
 3060 3065 3070
 Leu Asn Gly Asn Pro Thr Val Gln Asn Ile Asn Asp Lys Val Thr Glu
 3075 3080 3085
 10 Ala Asn Gln Ala Lys Asp Gln Leu Asn Thr Ala Arg Gln Gly Leu Thr
 3090 3095 3100
 Leu Asp Arg Gln Pro Ala Leu Thr Thr Leu His Gly Ala Ser Asn Leu
 3105 3110 3115 3120
 15 Asn Gln Ala Gln Gln Asn Asn Phe Thr Gln Gln Ile Asn Ala Ala Gln
 3125 3130 3135
 Asn His Ala Ala Leu Glu Thr Ile Lys Ser Asn Ile Thr Ala Leu Asn
 3140 3145 3150
 20 Thr Ala Met Thr Lys Leu Lys Asp Ser Val Ala Asp Asn Asn Thr Ile
 3155 3160 3165
 Lys Ser Asp Gln Asn Tyr Thr Asp Ala Thr Pro Ala Asn Lys Gln Ala
 3170 3175 3180
 25 Tyr Asp Asn Ala Val Asn Ala Ala Lys Gly Val Ile Gly Glu Thr Thr
 3185 3190 3195 3200
 Asn Pro Thr Met Asp Val Asn Thr Val Asn Gln Lys Ala Ala Ser Val
 3205 3210 3215
 30 Lys Ser Thr Lys Asp Ala Leu Asp Gly Gln Gln Asn Leu Gln Arg Ala
 3220 3225 3230
 Lys Thr Glu Ala Thr Asn Ala Ile Thr His Ala Ser Asp Leu Asn Gln
 3235 3240 3245
 35 Ala Gln Lys Asn Ala Leu Thr Gln Gln Val Asn Ser Ala Gln Asn Val
 3250 3255 3260
 Gln Ala Val Asn Asp Ile Lys Gln Thr Thr Gln Ser Leu Asn Thr Ala
 3265 3270 3275 3280
 40 Met Thr Gly Leu Lys Arg Gly Val Ala Asn His Asn Gln Val Val Gln
 3285 3290 3295
 Ser Asp Asn Tyr Val Asn Ala Asp Thr Asn Lys Lys Asn Asp Tyr Asn
 3300 3305 3310
 45 Asn Ala Tyr Asn His Ala Asn Asp Ile Ile Asn Gly Asn Ala Gln His
 3315 3320 3325
 Pro Val Ile Thr Pro Ser Asp Val Asn Asn Ala Leu Ser Asn Val Thr
 3330 3335 3340
 50 Ser Lys Glu His Ala Leu Asn Gly Glu Ala Lys Leu Asn Ala Ala Lys
 3345 3350 3355 3360
 Gln Glu Ala Asn Thr Ala Leu Gly His Leu Asn Asn Leu Asn Asn Ala
 3365 3370 3375
 55 Gln Arg Gln Asn Leu Gln Ser Gln Ile Asn Gly Ala His Gln Ile Asp
 3380 3385 3390
 Ala Val Asn Thr Ile Lys Gln Asn Ala Thr Asn Leu Asn Ser Ala Met
 3395 3400 3405

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Gly Asn Leu Arg Gln Ala Val Ala Asp Lys Asp Gln Val Lys Arg Thr
 3410 3415 3420
 5
 Glu Asp Tyr Ala Asp Ala Asp Thr Ala Lys Gln Asn Ala Tyr Asn Ser
 3425 3430 3435 3440
 Ala Val Ser Ser Ala Glu Thr Ile Ile Asn Gln Thr Thr Asn Pro Thr
 3445 3450 3455
 10
 Met Ser Val Asp Asp Val Asn Arg Ala Thr Ser Ala Val Thr Ser Asn
 3460 3465 3470
 Lys Asn Ala Leu Asn Gly Tyr Glu Lys Leu Ala Gln Ser Lys Thr Asp
 3475 3480 3485
 15
 Ala Ala Arg Ala Ile Asp Ala Leu Pro His Leu Asn Asn Ala Gln Lys
 3490 3495 3500
 Ala Asp Val Lys Ser Lys Ile Asn Ala Ala Ser Asn Ile Ala Gly Val
 3505 3510 3515 3520
 20
 Asn Thr Val Lys Gln Gln Gly Thr Asp Leu Asn Thr Ala Met Gly Asn
 3525 3530 3535
 Leu Gln Gly Ala Ile Asn Asp Glu Gln Thr Thr Leu Asn Ser Gln Asn
 3540 3545 3550
 Tyr Gln Asp Ala Thr Pro Ser Lys Lys Thr Ala Tyr Thr Asn Ala Val
 3555 3560 3565
 25
 Gln Ala Ala Lys Asp Ile Leu Asn Lys Ser Asn Gly Gln Asn Lys Thr
 3570 3575 3580
 Lys Asp Gln Val Thr Glu Ala Met Asn Gln Val Asn Ser Ala Lys Asn
 3585 3590 3595 3600
 30
 Asn Leu Asp Gly Thr Arg Leu Leu Asp Gln Ala Lys Gln Thr Ala Lys
 3605 3610 3615
 Gln Gln Leu Asn Asn Met Thr His Leu Thr Thr Ala Gln Lys Thr Asn
 3620 3625 3630
 35
 Leu Thr Asn Gln Ile Asn Ser Gly Thr Thr Val Ala Gly Val Gln Thr
 3635 3640 3645
 Val Gln Ser Asn Ala Asn Thr Leu Asp Gln Ala Met Asn Thr Leu Arg
 3650 3655 3660
 40
 Gln Ser Ile Ala Asn Lys Asp Ala Thr Lys Ala Ser Glu Asp Tyr Val
 3665 3670 3675 3680
 Asp Ala Asn Asn Asp Lys Gln Thr Ala Tyr Asn Asn Ala Val Ala Ala
 3685 3690 3695
 45
 Ala Glu Thr Ile Ile Asn Ala Asn Ser Asn Pro Glu Met Asn Pro Ser
 3700 3705 3710
 Thr Ile Thr Gln Lys Ala Glu Gln Val Asn Ser Ser Lys Thr Ala Leu
 3715 3720 3725
 50
 Asn Gly Asp Glu Asn Leu Ala Ala Ala Lys Gln Asn Ala Lys Thr Tyr
 3730 3735 3740
 Leu Asn Thr Leu Thr Ser Ile Thr Asp Ala Gln Lys Asn Asn Leu Ile
 3745 3750 3755 3760
 55
 Ser Gln Ile Thr Ser Ala Thr Arg Val Ser Gly Val Asp Thr Val Lys
 3765 3770 3775
 Gln Asn Ala Gln His Leu Asp Gln Ala Met Ala Ser Leu Gln Asn Gly

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	3780						3785					3790				
	Ile	Asn	Asn	Glu	Ser	Gln	Val	Lys	Ser	Ser	Glu	Lys	Tyr	Arg	Asp	Ala
			3795					3800					3805			
5	Asp	Thr	Asn	Lys	Gln	Gln	Glu	Tyr	Asp	Asn	Ala	Ile	Thr	Ala	Ala	Lys
		3810					3815					3820				
	Ala	Ile	Leu	Asn	Lys	Ser	Thr	Gly	Pro	Asn	Thr	Ala	Gln	Asn	Ala	Val
	3825					3830					3835					3840
10	Glu	Ala	Ala	Leu	Gln	Arg	Val	Asn	Asn	Ala	Lys	Asp	Ala	Leu	Asn	Gly
					3845					3850					3855	
	Asp	Ala	Lys	Leu	Ile	Ala	Ala	Gln	Asn	Ala	Ala	Lys	Gln	His	Leu	Gly
				3860					3865					3870		
15	Thr	Leu	Thr	His	Ile	Thr	Thr	Ala	Gln	Arg	Asn	Asp	Leu	Thr	Asn	Gln
			3875					3880					3885			
	Ile	Ser	Gln	Ala	Thr	Asn	Leu	Ala	Gly	Val	Glu	Ser	Val	Lys	Gln	Asn
		3890					3895					3900				
20	Ala	Asn	Ser	Leu	Asp	Gly	Ala	Met	Gly	Asn	Leu	Gln	Thr	Ala	Ile	Asn
	3905					3910					3915					3920
	Asp	Lys	Ser	Gly	Thr	Leu	Ala	Ser	Gln	Asn	Phe	Leu	Asp	Ala	Asp	Glu
					3925					3930					3935	
25	Gln	Lys	Arg	Asn	Ala	Tyr	Asn	Gln	Ala	Val	Ser	Ala	Ala	Glu	Thr	Ile
				3940					3945					3950		
	Leu	Asn	Lys	Gln	Thr	Gly	Pro	Asn	Thr	Ala	Lys	Thr	Ala	Val	Glu	Gln
			3955					3960						3965		
30	Ala	Leu	Asn	Asn	Val	Asn	Asn	Ala	Lys	His	Ala	Leu	Asn	Gly	Thr	Gln
	3970						3975					3980				
	Asn	Leu	Asn	Asn	Ala	Lys	Gln	Ala	Ala	Ile	Thr	Ala	Ile	Asn	Gly	Ala
	3985					3990					3995					4000
35	Ser	Asp	Leu	Asn	Gln	Lys	Gln	Lys	Asp	Ala	Leu	Lys	Ala	Gln	Ala	Asn
					4005					4010					4015	
	Gly	Ala	Gln	Arg	Val	Ser	Asn	Ala	Gln	Asp	Val	Gln	His	Asn	Ala	Thr
				4020					4025					4030		
40	Glu	Leu	Asn	Thr	Ala	Met	Gly	Thr	Leu	Lys	His	Ala	Ile	Ala	Asp	Lys
			4035					4040					4045			
	Thr	Asn	Thr	Leu	Ala	Ser	Ser	Lys	Tyr	Val	Asn	Ala	Asp	Ser	Thr	Lys
		4050					4055					4060				
45	Gln	Asn	Ala	Tyr	Thr	Thr	Lys	Val	Thr	Asn	Ala	Glu	His	Ile	Ile	Ser
	4065					4070					4075					4080
	Gly	Thr	Pro	Thr	Val	Val	Thr	Thr	Pro	Ser	Glu	Val	Thr	Ala	Ala	Ala
					4085					4090					4095	
50	Asn	Gln	Val	Asn	Ser	Ala	Lys	Gln	Glu	Leu	Asn	Gly	Asp	Glu	Arg	Leu
				4100					4105					4110		
	Arg	Glu	Ala	Lys	Gln	Asn	Ala	Asn	Thr	Ala	Ile	Asp	Ala	Leu	Thr	Gln
			4115					4120					4125			
55	Leu	Asn	Thr	Pro	Gln	Lys	Ala	Lys	Leu	Lys	Glu	Gln	Val	Gly	Gln	Ala
		4130					4135					4140				
	Asn	Arg	Leu	Glu	Asp	Val	Gln	Thr	Val	Gln	Thr	Asn	Gly	Gln	Ala	Leu
	4145					4150					4155					4160

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Asn Asn Ala Met Lys Gly Leu Arg Asp Ser Ile Ala Asn Glu Thr Thr
 4165 4170 4175
 Val Lys Thr Ser Gln Asn Tyr Thr Asp Ala Ser Pro Asn Asn Gln Ser
 5 4180 4185 4190
 Thr Tyr Asn Ser Ala Val Ser Asn Ala Lys Gly Ile Ile Asn Gln Thr
 4195 4200 4205
 Asn Asn Pro Thr Met Asp Thr Ser Ala Ile Thr Gln Ala Thr Thr Gln
 10 4210 4215 4220
 Val Asn Asn Ala Lys Asn Gly Leu Asn Gly Ala Glu Asn Leu Arg Asn
 4225 4230 4240
 Ala Gln Asn Thr Ala Lys Gln Asn Leu Asn Thr Leu Ser His Leu Thr
 15 4245 4250 4255
 Asn Asn Gln Lys Ser Ala Ile Ser Ser Gln Ile Asp Arg Ala Gly His
 4260 4265 4270
 Val Ser Glu Val Thr Ala Thr Lys Asn Ala Ala Thr Glu Leu Asn Thr
 20 4275 4280 4285
 Gln Met Gly Asn Leu Glu Gln Ala Ile His Asp Gln Asn Thr Val Lys
 4290 4295 4300
 Gln Ser Val Lys Phe Thr Asp Ala Asp Lys Ala Lys Arg Asp Ala Tyr
 25 4305 4310 4315 4320
 Thr Asn Ala Val Ser Arg Ala Glu Ala Ile Leu Asn Lys Thr Gln Gly
 4325 4330 4335
 Ala Asn Thr Ser Lys Gln Asp Val Glu Ala Ala Ile Gln Asn Val Ser
 30 4340 4345 4350
 Ser Ala Lys Asn Ala Leu Asn Gly Asp Gln Asn Val Thr Asn Ala Lys
 4355 4360 4365
 Asn Ala Ala Lys Asn Ala Leu Asn Asn Leu Thr Ser Ile Asn Asn Ala
 35 4370 4375 4380
 Gln Lys Arg Asp Leu Thr Thr Lys Ile Asp Gln Ala Thr Thr Val Ala
 4385 4390 4400
 Gly Val Glu Ala Val Ser Asn Thr Ser Thr Gln Leu Asn Thr Ala Met
 40 4405 4410 4415
 Ala Asn Leu Gln Asn Gly Ile Asn Asp Lys Lys Thr Asn Thr Leu Ala Ser
 4420 4425 4430
 Glu Asn Tyr His Asp Ala Asp Ser Asp Lys Lys Thr Ala Tyr Thr Gln
 45 4435 4440 4445
 Ala Val Thr Asn Ala Glu Asn Ile Leu Asn Lys Asn Ser Gly Ser Asn
 4450 4455 4460
 Leu Asp Lys Thr Ala Val Glu Asn Ala Leu Ser Gln Val Ala Asn Ala
 50 4465 4470 4475 4480
 Lys Gly Ala Leu Asn Gly Asn His Asn Leu Glu Gln Ala Lys Ser Asn
 4485 4490 4495
 Ala Asn Thr Thr Ile Asn Gly Leu Gln His Leu Thr Thr Ala Gln Lys
 55 4500 4505 4510
 Asp Lys Leu Lys Gln Gln Val Gln Gln Ala Gln Asn Val Ala Gly Val
 4515 4520 4525

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Asp Thr Val Lys Ser Ser Ala Asn Thr Leu Asn Gly Ala Met Gly Thr
 4530 4535 4540
 5 Leu Arg Asn Ser Ile Gln Asp Asn Thr Ala Thr Lys Asn Gly Gln Asn
 4545 4550 4555
 Tyr Leu Asp Ala Thr Glu Arg Asn Lys Thr Asn Tyr Asn Asn Ala Val
 4565 4570 4575
 10 Asp Ser Ala Asn Gly Val Ile Asn Ala Thr Ser Asn Pro Asn Met Asp
 4580 4585 4590
 Ala Asn Ala Ile Asn Gln Ile Ala Thr Gln Val Thr Ser Thr Lys Asn
 4595 4600 4605
 15 Ala Leu Asp Gly Thr His Asn Leu Thr Gln Ala Lys Gln Thr Ala Thr
 4610 4615 4620
 Asn Ala Ile Asp Gly Ala Thr Asn Leu Asn Lys Ala Gln Lys Asp Ala
 4625 4630 4635 4640
 20 Leu Lys Ala Gln Val Thr Ser Ala Gln Arg Val Ala Asn Val Thr Ser
 4645 4650 4655
 Ile Gln Gln Thr Ala Asn Glu Leu Asn Thr Ala Met Gly Gln Leu Gln
 4660 4665 4670
 His Gly Ile Asp Asp Glu Asn Ala Thr Lys Gln Thr Gln Lys Tyr Arg
 4675 4680 4685
 25 Asp Ala Glu Gln Ser Lys Lys Thr Ala Tyr Asp Gln Ala Val Ala Ala
 4690 4695 4700
 Ala Lys Ala Ile Leu Asn Lys Gln Thr Gly Ser Asn Ser Asp Lys Ala
 4705 4710 4715 4720
 30 Ala Val Asp Arg Ala Leu Gln Gln Val Thr Ser Thr Lys Asp Ala Leu
 4725 4730 4735
 Asn Gly Asp Ala Lys Leu Ala Glu Ala Lys Ala Ala Lys Gln Asn
 4740 4745 4750
 35 Leu Gly Thr Leu Asn His Ile Thr Asn Ala Gln Arg Thr Asp Leu Glu
 4755 4760 4765
 Gly Gln Ile Asn Gln Ala Thr Thr Val Asp Gly Val Asn Thr Val Lys
 4770 4775 4780
 40 Thr Asn Ala Asn Thr Leu Asp Gly Ala Met Asn Ser Leu Gln Gly Ser
 4785 4790 4795 4800
 Ile Asn Asp Lys Asp Ala Thr Leu Arg Asn Gln Asn Tyr Leu Asp Ala
 4805 4810 4815
 45 Asp Glu Ser Lys Arg Asn Ala Tyr Thr Gln Ala Val Thr Ala Ala Glu
 4820 4825 4830
 Gly Ile Leu Asn Lys Gln Thr Gly Gly Asn Thr Ser Lys Ala Asp Val
 4835 4840 4845
 50 Asp Asn Ala Leu Asn Ala Val Thr Arg Ala Lys Ala Ala Leu Asn Gly
 4850 4855 4860
 Ala Asp Asn Leu Arg Asn Ala Lys Thr Ser Ala Thr Asn Thr Ile Asp
 4865 4870 4875 4880
 55 Gly Leu Pro Asn Leu Thr Gln Leu Gln Lys Asp Asn Leu Lys His Gln
 4885 4890 4895
 Val Glu Gln Ala Gln Asn Val Ala Gly Val Asn Gly Val Lys Asp Lys

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	4900	4905	4910
	Gly Asn Thr Leu Asn Thr Ala Met Gly Ala Leu Arg Thr Ser Ile Gln 4915 4920 4925		
5	Asn Asp Asn Thr Thr Lys Thr Ser Gln Asn Tyr Leu Asp Ala Ser Asp 4930 4935 4940		
	Ser Asn Lys Asn Asn Tyr Asn Thr Ala Val Asn Asn Ala Asn Gly Val 4945 4950 4955 4960		
10	Ile Asn Ala Thr Asn Asn Pro Asn Met Asp Ala Asn Ala Ile Asn Gly 4965 4970 4975		
	Met Ala Asn Gln Val Asn Thr Thr Lys Ala Ala Leu Asn Gly Ala Gln 4980 4985 4990		
15	Asn Leu Ala Gln Ala Lys Thr Asn Ala Thr Asn Thr Ile Asn Asn Ala 4995 5000 5005		
	His Asp Leu Asn Gln Lys Gln Lys Asp Ala Leu Lys Thr Gln Val Asn 5010 5015 5020		
20	Asn Ala Gln Arg Val Ser Asp Ala Asn Asn Val Gln His Thr Ala Thr 5025 5030 5035 5040		
	Glu Leu Asn Ser Ala Met Thr Ala Leu Lys Ala Ala Ile Ala Asp Lys 5045 5050 5055		
25	Glu Arg Thr Lys Ala Ser Gly Asn Tyr Val Asn Ala Asp Gln Glu Lys 5060 5065 5070		
	Arg Gln Ala Tyr Asp Ser Lys Val Thr Asn Ala Glu Asn Ile Ile Ser 5075 5080 5085		
30	Gly Thr Pro Asn Ala Thr Leu Thr Val Asn Asp Val Asn Ser Ala Ala 5090 5095 5100		
	Ser Gln Val Asn Ala Ala Lys Thr Ala Leu Asn Gly Asp Asn Asn Leu 5105 5110 5115 5120		
35	Arg Val Ala Lys Glu His Ala Asn Asn Thr Ile Asp Gly Leu Ala Gln 5125 5130 5135		
	Leu Asn Asn Ala Gln Lys Ala Lys Leu Lys Glu Gln Val Gln Ser Ala 5140 5145 5150		
40	Thr Thr Leu Asp Gly Val Gln Thr Val Lys Asn Ser Ser Gln Thr Leu 5155 5160 5165		
	Asn Thr Ala Met Lys Gly Leu Arg Asp Ser Ile Ala Asn Glu Ala Thr 5170 5175 5180		
45	Ile Lys Ala Gly Gln Asn Tyr Thr Asp Ala Ser Pro Asn Asn Arg Asn 5185 5190 5195 5200		
	Glu Tyr Asp Ser Ala Val Thr Ala Ala Lys Ala Ile Ile Asn Gln Thr 5205 5210 5215		
50	Ser Asn Pro Thr Met Glu Pro Asn Thr Ile Thr Gln Val Thr Ser Gln 5220 5225 5230		
	Val Thr Thr Lys Glu Gln Ala Leu Asn Gly Ala Arg Asn Leu Ala Gln 5235 5240 5245		
55	Ala Lys Thr Thr Ala Lys Asn Asn Leu Asn Asn Leu Thr Ser Ile Asn 5250 5255 5260		
	Asn Ala Gln Lys Asp Ala Leu Thr Arg Ser Ile Asp Gly Ala Thr Thr 5265 5270 5275 5280		

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Val Ala Gly Val Asn Gln Glu Thr Ala Lys Ala Thr Glu Leu Asn Asn
 5285 5290 5295
 Ala Met His Ser Leu Gln Asn Gly Ile Asn Asp Glu Thr Gln Thr Lys
 5300 5305 5310
 Gln Thr Gln Lys Tyr Leu Asp Ala Glu Pro Ser Lys Lys Ser Ala Tyr
 5315 5320 5325
 Asp Gln Ala Val Asn Ala Ala Lys Ala Ile Leu Thr Lys Ala Ser Gly
 5330 5335 5340
 Gln Asn Val Asp Lys Ala Ala Val Glu Gln Ala Leu Gln Asn Val Asn
 5345 5350 5355 5360
 Ser Thr Lys Thr Ala Leu Asn Gly Asp Ala Lys Leu Asn Glu Ala Lys
 5365 5370 5375
 Ala Ala Ala Lys Gln Thr Leu Gly Thr Leu Thr His Ile Asn Asn Ala
 5380 5385 5390
 Gln Arg Thr Ala Leu Asp Asn Glu Ile Thr Gln Ala Thr Asn Val Glu
 5395 5400 5405
 Gly Val Asn Thr Val Lys Ala Lys Ala Gln Gln Leu Asp Gly Ala Met
 5410 5415 5420
 Gly Gln Leu Glu Thr Ser Ile Arg Asp Lys Asp Thr Thr Leu Gln Ser
 5425 5430 5435 5440
 Gln Asn Tyr Gln Asp Ala Asp Asp Ala Lys Arg Thr Ala Tyr Ser Gln
 5445 5450 5455
 Ala Val Asn Ala Ala Ala Thr Ile Leu Asn Lys Thr Ala Gly Gly Asn
 5460 5465 5470
 Thr Pro Lys Ala Asp Val Glu Arg Ala Met Gln Ala Val Thr Gln Ala
 5475 5480 5485
 Asn Thr Ala Leu Asn Gly Ile Gln Asn Leu Asp Arg Ala Lys Gln Ala
 5490 5495 5500
 Ala Asn Thr Ala Ile Thr Asn Ala Ser Asp Leu Asn Thr Lys Gln Lys
 5505 5510 5515 5520
 Glu Ala Leu Lys Ala Gln Val Thr Ser Ala Gly Arg Val Ser Ala Ala
 5525 5530 5535
 Asn Gly Val Glu His Thr Ala Thr Glu Leu Asn Thr Ala Met Thr Ala
 5540 5545 5550
 Leu Lys Arg Ala Ile Ala Asp Lys Ala Glu Thr Lys Ala Ser Gly Asn
 5555 5560 5565
 Tyr Val Asn Ala Asp Ala Asn Lys Arg Gln Ala Tyr Asp Glu Lys Val
 5570 5575 5580
 Thr Ala Ala Glu Asn Ile Val Ser Gly Thr Pro Thr Pro Thr Leu Thr
 5585 5590 5595 5600
 Pro Ala Asp Val Thr Asn Ala Ala Thr Gln Val Thr Asn Ala Lys Thr
 5605 5610 5615
 Gln Leu Asn Gly Asn His Asn Leu Glu Val Ala Lys Gln Asn Ala Asn
 5620 5625 5630
 Thr Ala Ile Asp Gly Leu Thr Ser Leu Asn Gly Pro Gln Lys Ala Lys
 5635 5640 5645

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Leu Lys Glu Gln Val Gly Gln Ala Thr Thr Leu Pro Asn Val Gln Thr
 5650 5655 5660
 Val Arg Asp Asn Ala Gln Thr Leu Asn Thr Ala Met Lys Gly Leu Arg
 5665 5670 5675 5680
 5 Asp Ser Ile Ala Asn Glu Ala Thr Ile Lys Ala Gly Gln Asn Tyr Thr
 5685 5690 5695
 Asp Ala Ser Gln Asn Lys Gln Thr Asp Tyr Asn Ser Ala Val Thr Ala
 5700 5705 5710
 10 Ala Lys Ala Ile Ile Gly Gln Thr Thr Ser Pro Ser Met Asn Ala Gln
 5715 5720 5725
 Glu Ile Asn Gln Ala Lys Asp Gln Val Thr Ala Lys Gln Gln Ala Leu
 5730 5735 5740
 15 Asn Gly Gln Glu Asn Leu Arg Thr Ala Gln Thr Asn Ala Lys Gln His
 5745 5750 5755 5760
 Leu Asn Gly Leu Ser Asp Leu Thr Asp Ala Gln Lys Asp Ala Val Lys
 5765 5770 5775
 20 Arg Gln Ile Glu Gly Ala Thr His Val Asn Glu Val Thr Gln Ala Gln
 5780 5785 5790
 Asn Asn Ala Asp Ala Leu Asn Thr Ala Met Thr Asn Leu Lys Asn Gly
 5795 5800 5805
 25 Ile Gln Asp Gln Asn Thr Ile Lys Gln Gly Val Asn Phe Thr Asp Ala
 5810 5815 5820
 Asp Glu Ala Lys Arg Asn Ala Tyr Thr Asn Ala Val Thr Gln Ala Glu
 5825 5830 5835 5840
 30 Gln Ile Leu Asn Lys Ala Gln Gly Pro Asn Thr Ser Lys Asp Gly Val
 5845 5850 5855
 Glu Thr Ala Leu Glu Asn Val Gln Arg Ala Lys Asn Glu Leu Asn Gly
 5860 5865 5870
 35 Asn Gln Asn Val Ala Asn Ala Lys Thr Thr Ala Lys Asn Ala Leu Asn
 5875 5880 5885
 Asn Leu Thr Ser Ile Asn Asn Ala Gln Lys Glu Ala Leu Lys Ser Gln
 5890 5895 5900
 40 Ile Glu Gly Ala Thr Thr Val Ala Gly Val Asn Gln Val Ser Thr Thr
 5905 5910 5915 5920
 Ala Ser Glu Leu Asn Thr Ala Met Ser Asn Leu Gln Asn Gly Ile Asn
 5925 5930 5935
 45 Asp Glu Ala Ala Thr Lys Ala Ala Leu Asn Gly Thr Gln Asn Leu Glu
 5940 5945 5950
 Lys Ala Lys Gln His Ala Asn Thr Ala Ile Asp Gly Leu Ser His Leu
 5955 5960 5965
 50 Thr Asn Ala Gln Lys Glu Ala Leu Lys Gln Leu Val Gln Gln Ser Thr
 5970 5975 5980
 Thr Val Ala Glu Ala Gln Gly Asn Glu Gln Lys Ala Asn Asn Val Asp
 5985 5990 5995 6000
 55 Ala Ala Met Asp Lys Leu Arg Gln Ser Ile Ala Asp Asn Ala Thr Thr
 6005 6010 6015
 Lys Gln Asn Gln Asn Tyr Thr Asp Ala Ser Gln Asn Lys Lys Asp Ala

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		6020				6025				6030							
		Tyr	Asn	Asn	Ala	Val	Thr	Thr	Ala	Gln	Gly	Ile	Ile	Asp	Gln	Thr	Thr
				6035					6040					6045			
5		Ser	Pro	Thr	Leu	Asp	Pro	Thr	Val	Ile	Asn	Gln	Ala	Ala	Gly	Gln	Val
			6050					6055					6060				
		Ser	Thr	Thr	Lys	Asn	Ala	Leu	Asn	Gly	Asn	Glu	Asn	Leu	Glu	Ala	Ala
		6065				6070				6075							6080
10		Lys	Gln	Gln	Ala	Ser	Gln	Ser	Leu	Gly	Ser	Leu	Asp	Asn	Leu	Asn	Asn
						6085					6090					6095	
		Ala	Gln	Lys	Gln	Thr	Val	Thr	Asp	Gln	Ile	Asn	Gly	Ala	His	Thr	Val
					6100					6105					6110		
15		Asp	Glu	Ala	Asn	Gln	Ile	Lys	Gln	Asn	Ala	Gln	Asn	Leu	Asn	Thr	Ala
			6115						6120					6125			
		Met	Gly	Asn	Leu	Lys	Gln	Ala	Ile	Ala	Asp	Lys	Asp	Ala	Thr	Lys	Ala
		6130						6135					6140				
20		Thr	Val	Asn	Phe	Thr	Asp	Ala	Asp	Gln	Ala	Lys	Gln	Gln	Ala	Tyr	Asn
		6145					6150					6155					6160
		Thr	Ala	Val	Thr	Asn	Ala	Glu	Asn	Ile	Ile	Ser	Lys	Ala	Asn	Gly	Gly
						6165					6170					6175	
25		Asn	Ala	Thr	Gln	Ala	Glu	Val	Glu	Gln	Ala	Ile	Lys	Gln	Val	Asn	Ala
					6180					6185					6190		
		Ala	Lys	Gln	Ala	Leu	Asn	Gly	Asn	Ala	Asn	Val	Gln	His	Ala	Lys	Asp
				6195					6200					6205			
30		Glu	Ala	Thr	Ala	Leu	Ile	Asn	Ser	Ser	Asn	Asp	Leu	Asn	Gln	Ala	Gln
		6210						6215					6220				
		Lys	Asp	Ala	Leu	Lys	Gln	Gln	Val	Gln	Asn	Ala	Thr	Thr	Val	Ala	Gly
		6225					6230				6235						6240
35		Val	Asn	Asn	Val	Lys	Gln	Thr	Ala	Gln	Glu	Leu	Asn	Asn	Ala	Met	Thr
						6245					6250					6255	
		Gln	Leu	Lys	Gln	Gly	Ile	Ala	Asp	Lys	Glu	Gln	Thr	Lys	Ala	Asp	Gly
					6260					6265					6270		
40		Asn	Phe	Val	Asn	Ala	Asp	Pro	Asp	Lys	Gln	Asn	Ala	Tyr	Asn	Gln	Ala
				6275					6280					6285			
		Val	Ala	Lys	Ala	Glu	Ala	Leu	Ile	Ser	Ala	Thr	Pro	Asp	Val	Val	Val
				6290				6295					6300				
45		Thr	Pro	Ser	Glu	Ile	Thr	Ala	Ala	Leu	Asn	Lys	Val	Thr	Gln	Ala	Lys
		6305					6310					6315					6320
		Asn	Asp	Leu	Asn	Gly	Asn	Thr	Asn	Leu	Ala	Thr	Ala	Lys	Gln	Asn	Val
						6325					6330					6335	
50		Gln	His	Ala	Ile	Asp	Gln	Leu	Pro	Asn	Leu	Asn	Gln	Ala	Gln	Arg	Asp
					6340					6345					6350		
55		Glu	Tyr	Ser	Lys	Gln	Ile	Thr	Gln	Ala	Thr	Leu	Val	Pro	Asn	Val	Asn
				6355					6360					6365			
		Ala	Ile	Gln	Gln	Ala	Ala	Thr	Thr	Leu	Asn	Asp	Ala	Met	Thr	Gln	Leu
			6370					6375					6380				
60		Lys	Gln	Gly	Ile	Ala	Asn	Lys	Ala	Gln	Ile	Lys	Gly	Ser	Glu	Asn	Tyr
		6385					6390					6395					6400

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His Asp Ala Asp Thr Asp Lys Gln Thr Ala Tyr Asp Asn Ala Val Thr
 6405 6410 6415
 5 Lys Ala Glu Glu Leu Leu Lys Gln Thr Thr Asn Pro Thr Met Asp Pro
 6420 6425 6430
 Asn Thr Ile Gln Gln Ala Leu Thr Lys Val Asn Asp Thr Asn Gln Ala
 6435 6440 6445
 10 Leu Asn Gly Asn Gln Lys Leu Ala Asp Ala Lys Gln Asp Ala Lys Thr
 6450 6455 6460
 Thr Leu Gly Thr Leu Asp His Leu Asn Asp Ala Gln Lys Gln Ala Leu
 6465 6470 6475 6480
 15 Thr Thr Gln Val Glu Gln Ala Pro Asp Ile Ala Thr Val Asn Asn Val
 6485 6490 6495
 Lys Gln Asn Ala Gln Asn Leu Asn Asn Ala Met Thr Asn Leu Asn Asn
 6500 6505 6510
 20 Ala Leu Gln Asp Lys Thr Glu Thr Leu Asn Ser Ile Asn Phe Thr Asp
 6515 6520 6525
 Ala Asp Gln Ala Lys Lys Asp Ala Tyr Thr Asn Ala Val Ser His Ala
 6530 6535 6540
 25 Glu Gly Ile Leu Ser Lys Ala Asn Gly Ser Asn Ala Ser Gln Thr Glu
 6545 6550 6555 6560
 Val Glu Gln Ala Met Gln Arg Val Asn Glu Ala Lys Gln Ala Leu Asn
 6565 6570 6575
 30 Gly Asn Asp Asn Val Gln Arg Ala Lys Asp Ala Ala Lys Gln Val Ile
 6580 6585 6590
 Thr Asn Ala Asn Asp Leu Asn Gln Ala Met Thr Gln Leu Lys Gln Gly
 6595 6600 6605
 35 Ile Ala Asp Lys Asp Gln Thr Lys Ala Asn Gly Asn Phe Val Asn Ala
 6610 6615 6620
 Asp Thr Asp Lys Gln Asn Ala Tyr Asn Asn Ala Val Ala His Ala Glu
 6625 6630 6635 6640
 40 Gln Ile Ile Ser Gly Thr Pro Asn Ala Asn Val Asp Pro Gln Gln Val
 6645 6650 6655
 Ala Gln Ala Leu Gln Gln Val Asn Gln Ala Lys Gly Asp Leu Asn Gly
 6660 6665 6670
 45 Asn His Asn Leu Gln Val Ala Lys Asp Asn Ala Asn Thr Ala Ile Asp
 6675 6680 6685
 Gln Leu Pro Asn Leu Asn Gln Pro Gln Lys Thr Ala Leu Lys Asp Gln
 6690 6695 6700
 50 Val Ser His Ala Glu Leu Val Thr Gly Val Asn Ala Ile Lys Gln Asn
 6705 6710 6715 6720
 Ala Asp Ala Leu Asn Asn Ala Met Gly Thr Leu Lys Gln Gln Ile Gln
 6725 6730 6735
 55 Ala Asn Ser Gln Val Pro Gln Ser Val Asp Phe Thr Gln Ala Asp Gln
 6740 6745 6750
 Asp Lys Gln Gln Ala Tyr Asn Asn Ala Ala Asn Gln Ala Gln Gln Ile
 6755 6760 6765

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Ala Asn Gly Ile Pro Thr Pro Val Leu Thr Pro Asp Thr Val Thr Gln
6770 6775 6780

Ala Val Thr Thr Met Asn Gln Ala Lys Asp Ala Leu Asn Gly Asp Glu
5 6785 6790 6795 6800

Lys Leu Ala Gln Ala Lys Gln Glu Ala Leu Ala Asn Leu Asp Thr Leu
6805 6810 6815

Arg Asp Leu Asn Gln Pro Gln Arg Asp Ala Leu Arg Asn Gln Ile Asn
10 6820 6825 6830

Gln Ala Gln Ala Leu Ala Thr Val Glu Gln Thr Lys Gln Asn Ala Gln
6835 6840 6845

Asn Val Asn Thr Ala Met Ser Asn Leu Lys Gln Gly Ile Ala Asn Lys
15 6850 6855 6860

Asp Thr Val Lys Ala Ser Glu Asn Tyr His Asp Ala Asp Ala Asp Lys
6865 6870 6875 6880

Gln Thr Ala Tyr Thr Asn Ala Val Ser Gln Ala Glu Gly Ile Ile Asn
20 6885 6890 6895

Gln Thr Thr Asn Pro Thr Leu Asn Pro Asp Glu Ile Thr Arg Ala Leu
6900 6905 6910

Thr Gln Val Thr Asp Ala Lys Asn Gly Leu Asn Gly Glu Ala Lys Leu
6915 6920 6925

Ala Thr Glu Lys Gln Asn Ala Lys Asp Ala Val Ser Gly Met Thr His
25 6930 6935 6940

Leu Asn Asp Ala Gln Lys Gln Ala Leu Lys Gly Gln Ile Asp Gln Ser
6945 6950 6955 6960

Pro Glu Ile Ala Thr Val Asn Gln Val Lys Gln Thr Ala Thr Ser Leu
30 6965 6970 6975

Asp Gln Ala Met Asp Gln Leu Ser Gln Ala Ile Asn Asp Lys Ala Gln
6980 6985 6990

Thr Leu Ala Asp Gly Asn Tyr Leu Asn Ala Asp Pro Asp Lys Gln Asn
35 6995 7000 7005

Ala Tyr Lys Gln Ala Val Ala Lys Ala Glu Ala Leu Leu Asn Lys Gln
7010 7015 7020

Ser Gly Thr Asn Glu Val Gln Ala Gln Val Glu Ser Ile Thr Asn Glu
40 7025 7030 7035 7040

Val Asn Ala Ala Lys Gln Ala Leu Asn Gly Asn Asp Asn Leu Ala Asn
7045 7050 7055

Ala Lys Gln Gln Ala Lys Gln Gln Leu Ala Asn Leu Thr His Leu Asn
45 7060 7065 7070

Asp Ala Gln Lys Gln Ser Phe Glu Ser Gln Ile Thr Gln Ala Pro Leu
7075 7080 7085

Val Thr Asp Val Thr Thr Ile Asn Gln Lys Ala Gln Thr Leu Asp His
50 7090 7095 7100

Ala Met Glu Leu Leu Arg Asn Ser Val Ala Asp Asn Gln Thr Thr Leu
7105 7110 7115 7120

Ala Ser Glu Asp Tyr His Asp Ala Thr Ala Gln Arg Gln Asn Asp Tyr
55 7125 7130 7135

Asn Gln Ala Val Thr Ala Ala Asn Asn Ile Ile Asn Gln Thr Thr Ser

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			7140					7145					7150			
	Pro	Thr	Met	Asn	Pro	Asp	Asp	Val	Asn	Gly	Ala	Thr	Thr	Gln	Val	Asn
			7155					7160					7165			
5	Asn	Thr	Lys	Val	Ala	Leu	Asp	Gly	Asp	Glu	Asn	Leu	Ala	Ala	Ala	Lys
			7170					7175					7180			
	Gln	Gln	Ala	Asn	Asn	Arg	Leu	Asp	Gln	Leu	Asp	His	Leu	Asn	Asn	Ala
			7185					7190					7195			7200
10	Gln	Lys	Gln	Gln	Leu	Gln	Ser	Gln	Ile	Thr	Gln	Ser	Ser	Asp	Ile	Ala
					7205					7210					7215	
	Ala	Val	Asn	Gly	His	Lys	Gln	Thr	Ala	Glu	Ser	Leu	Asn	Thr	Ala	Met
				7220					7225					7230		
15	Gly	Asn	Leu	Ile	Asn	Ala	Ile	Ala	Asp	His	Gln	Ala	Val	Glu	Gln	Arg
			7235						7240					7245		
	Gly	Asn	Phe	Ile	Asn	Ala	Asp	Thr	Asp	Lys	Gln	Thr	Ala	Tyr	Asn	Thr
			7250					7255						7260		
20	Ala	Val	Asn	Glu	Ala	Ala	Ala	Met	Ile	Asn	Lys	Gln	Thr	Gly	Gln	Asn
						7270						7275				7280
	Ala	Asn	Gln	Thr	Glu	Val	Glu	Gln	Ala	Ile	Thr	Lys	Val	Gln	Thr	Thr
					7285										7295	
25	Leu	Gln	Ala	Leu	Asn	Gly	Asp	His	Asn	Leu	Gln	Val	Ala	Lys	Thr	Asn
				7300						7305					7310	
	Ala	Thr	Gln	Ala	Ile	Asp	Ala	Leu	Thr	Ser	Leu	Asn	Asp	Pro	Gln	Lys
				7315					7320					7325		
30	Thr	Ala	Leu	Lys	Asp	Gln	Val	Thr	Ala	Ala	Thr	Leu	Val	Thr	Ala	Val
							7335							7340		
	His	Gln	Ile	Glu	Gln	Asn	Ala	Asn	Thr	Leu	Asn	Gln	Ala	Met	His	Gly
						7350										7360
35	Leu	Arg	Gln	Ser	Ile	Gln	Asp	Asn	Ala	Ala	Thr	Lys	Ala	Asn	Ser	Lys
					7365											7375
	Tyr	Ile	Asn	Glu	Asp	Gln	Pro	Glu	Gln	Gln	Asn	Tyr	Asp	Gln	Ala	Val
				7380						7385					7390	
40	Gln	Ala	Ala	Asn	Asn	Ile	Ile	Asn	Glu	Gln	Thr	Ala	Thr	Leu	Asp	Asn
				7395						7400				7405		
	Asn	Ala	Ile	Asn	Gln	Ala	Ala	Thr	Thr	Val	Asn	Thr	Thr	Lys	Ala	Ala
							7415							7420		
45	Leu	His	Gly	Asp	Val	Lys	Leu	Gln	Asn	Asp	Lys	Asp	His	Ala	Lys	Gln
							7430					7435				7440
	Thr	Val	Ser	Gln	Leu	Ala	His	Leu	Asn	Asn	Ala	Gln	Lys	His	Met	Glu
					7445						7450				7455	
50	Asp	Thr	Leu	Ile	Asp	Ser	Glu	Thr	Thr	Arg	Thr	Ala	Val	Lys	Gln	Asp
				7460							7465				7470	
	Leu	Thr	Glu	Ala	Gln	Ala	Leu	Asp	Gln	Leu	Met	Asp	Ala	Leu	Gln	Gln
								7480						7485		
55	Ser	Ile	Ala	Asp	Lys	Asp	Ala	Thr	Arg	Ala	Ser	Ser	Ala	Tyr	Val	Asn
								7495						7500		
	Ala	Glu	Pro	Asn	Lys	Lys	Gln	Ser	Tyr	Asp	Glu	Ala	Val	Gln	Asn	Ala
																7520
							7510									
											7515					

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Glu Ser Ile Ile Ala Gly Leu Asn Asn Pro Thr Ile Asn Lys Gly Asn
 7525 7530 7535
 Val Ser Ser Ala Thr Gln Ala Val Ile Ser Ser Lys Asn Ala Leu Asp
 7540 7545 7550
 Gly Val Glu Arg Leu Ala Gln Asp Lys Gln Thr Ala Gly Asn Ser Leu
 7555 7560 7565
 Asn His Leu Asp Gln Leu Thr Pro Ala Gln Gln Gln Ala Leu Glu Asn
 7570 7575 7580
 Gln Ile Asn Asn Ala Thr Thr Arg Asp Lys Val Ala Glu Ile Ile Ala
 7585 7590 7595 7600
 Gln Ala Gln Ala Leu Asn Glu Ala Met Lys Ala Leu Lys Glu Ser Ile
 7605 7610 7615
 Lys Asp Gln Pro Gln Thr Glu Ala Ser Ser Lys Phe Ile Asn Glu Asp
 7620 7625 7630
 Gln Ala Gln Lys Asp Ala Tyr Thr Gln Ala Val Gln His Ala Lys Asp
 7635 7640 7645
 Leu Ile Asn Lys Thr Thr Asp Pro Thr Leu Ala Lys Ser Ile Ile Asp
 7650 7655 7660
 Gln Ala Thr Gln Ala Val Thr Asp Ala Lys Asn Asn Leu His Gly Asp
 7665 7670 7675 7680
 Gln Lys Leu Ala Gln Asp Lys Gln Arg Ala Thr Glu Thr Leu Asn Asn
 7685 7690 7695
 Leu Ser Asn Leu Asn Thr Pro Gln Arg Gln Ala Leu Glu Asn Gln Ile
 7700 7705 7710
 Asn Asn Ala Ala Thr Arg Gly Glu Val Ala Gln Lys Leu Thr Glu Ala
 7715 7720 7725
 Gln Ala Leu Asn Gln Ala Met Glu Ala Leu Arg Asn Ser Ile Gln Asp
 7730 7735 7740
 Gln Gln Gln Thr Glu Ala Gly Ser Lys Phe Ile Asn Glu Asp Lys Pro
 7745 7750 7755 7760
 Gln Lys Asp Ala Tyr Gln Ala Ala Val Gln Asn Ala Lys Asp Leu Ile
 7765 7770 7775
 Asn Gln Thr Asn Asn Pro Thr Leu Asp Lys Ala Gln Val Glu Gln Leu
 7780 7785 7790
 Thr Gln Ala Val Asn Gln Ala Lys Asp Asn Leu His Gly Asp Gln Lys
 7795 7800 7805
 Leu Ala Asp Asp Lys Gln His Ala Val Thr Asp Leu Asn Gln Leu Asn
 7810 7815 7820
 Gly Leu Asn Asn Pro Gln Arg Gln Ala Leu Glu Ser Gln Ile Asn Asn
 7825 7830 7835 7840
 Ala Ala Thr Arg Gly Glu Val Ala Gln Lys Leu Ala Glu Ala Lys Ala
 7845 7850 7855
 Leu Asp Gln Ala Met Gln Ala Leu Arg Asn Ser Ile Gln Asp Gln Gln
 7860 7865 7870
 Gln Thr Glu Ser Gly Ser Lys Phe Ile Asn Glu Asp Lys Pro Gln Lys
 7875 7880 7885

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Asp Ala Tyr Gln Ala Ala Val Gln Asn Ala Lys Asp Leu Ile Asn Gln
 7890 7895 7900
 Thr Gly Asn Pro Thr Leu Asp Lys Ser Gln Val Glu Gln Leu Thr Gln
 5 7905 7910 7915 7920
 Ala Val Thr Thr Ala Lys Asp Asn Leu His Gly Asp Gln Lys Leu Ala
 7925 7930 7935
 Arg Asp Gln Gln Gln Ala Val Thr Thr Val Asn Ala Leu Pro Asn Leu
 10 7940 7945 7950
 Asn His Ala Gln Gln Gln Ala Leu Thr Asp Ala Ile Asn Ala Ala Pro
 7955 7960 7965
 Thr Arg Thr Glu Val Ala Gln His Val Gln Thr Ala Thr Glu Leu Asp
 15 7970 7975 7980
 His Ala Met Glu Thr Leu Lys Asn Lys Val Asp Gln Val Asn Thr Asp
 7985 7990 7995 8000
 Lys Ala Gln Pro Asn Tyr Thr Glu Ala Ser Thr Asp Lys Lys Glu Ala
 20 8005 8010 8015
 Val Asp Gln Ala Leu Gln Ala Ala Glu Ser Ile Thr Asp Pro Thr Asn
 8020 8025 8030
 Gly Ser Asn Ala Asn Lys Asp Ala Val Asp Gln Val Leu Thr Lys Leu
 8035 8040 8045
 25 Gln Glu Lys Glu Asn Glu Leu Asn Gly Asn Glu Arg Val Ala Glu Ala
 8050 8055 8060
 Lys Thr Gln Ala Lys Gln Thr Ile Asp Gln Leu Thr His Leu Asn Ala
 8065 8070 8075 8080
 30 Asp Gln Ile Ala Thr Ala Lys Gln Asn Ile Asp Gln Ala Thr Lys Leu
 8085 8090 8095
 Gln Pro Ile Ala Glu Leu Val Asp Gln Ala Thr Gln Leu Asn Gln Ser
 8100 8105 8110
 35 Met Asp Gln Leu Gln Gln Ala Val Asn Glu His Ala Asn Val Glu Gln
 8115 8120 8125
 Thr Val Asp Tyr Thr Gln Ala Asp Ser Asp Lys Gln Asn Ala Tyr Lys
 8130 8135 8140
 40 Gln Ala Ile Ala Asp Ala Glu Asn Val Leu Lys Gln Asn Ala Asn Lys
 8145 8150 8155 8160
 Gln Gln Val Asp Gln Ala Leu Gln Asn Ile Leu Asn Ala Lys Gln Ala
 8165 8170 8175
 45 Leu Asn Gly Asp Glu Arg Val Ala Leu Ala Lys Thr Asn Gly Lys His
 8180 8185 8190
 Asp Ile Asp Gln Leu Asn Ala Leu Asn Asn Ala Gln Gln Asp Gly Phe
 8195 8200 8205
 50 Lys Gly Arg Ile Asp Gln Ser Asn Asp Leu Asn Gln Ile Gln Gln Ile
 8210 8215 8220
 Val Asp Glu Ala Lys Ala Leu Asn Arg Ala Met Asp Gln Leu Ser Gln
 8225 8230 8235 8240
 55 Glu Ile Thr Asp Asn Glu Gly Arg Thr Lys Gly Ser Thr Asn Tyr Val
 8245 8250 8255
 Asn Ala Asp Thr Gln Val Lys Gln Val Tyr Asp Glu Thr Val Asp Lys

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	8260		8265		8270
	Ala Lys Gln Ala Leu Asp Lys Ser Thr Gly Gln Asn Leu Thr Ala Lys				
	8275		8280		8285
5	Gln Val Ile Lys Leu Asn Asp Ala Val Thr Ala Ala Lys Lys Ala Leu				
	8290		8295		8300
	Asn Gly Glu Glu Arg Leu Asn Asn Arg Lys Ala Glu Ala Leu Gln Arg				
	8305		8310		8315
10	Leu Asp Gln Leu Thr His Leu Asn Asn Ala Gln Arg Gln Leu Ala Ile				
			8325		8330
	Gln Gln Ile Asn Asn Ala Glu Thr Leu Asn Lys Ala Ser Arg Ala Ile				
			8340		8345
	Asn Arg Ala Thr Lys Leu Asp Asn Ala Met Gly Ser Val Gln Gln Tyr				
			8355		8360
15	Ile Asp Glu Gln His Leu Gly Val Ile Ser Ser Thr Asn Tyr Ile Asn				
			8370		8375
	Ala Asp Asp Asn Leu Lys Ala Asn Tyr Asp Asn Ala Ile Ala Asn Ala				
			8385		8390
20	Ala His Glu Leu Asp Lys Val Gln Gly Asn Ala Ile Ala Lys Ala Glu				
			8405		8410
	Ala Glu Gln Leu Lys Gln Asn Ile Ile Asp Ala Gln Asn Ala Leu Asn				
			8420		8425
25	Gly Asp Gln Asn Leu Ala Asn Ala Lys Asp Lys Ala Asn Ala Phe Val				
			8435		8440
	Asn Ser Leu Asn Gly Leu Asn Gln Gln Gln Gln Asp Leu Ala His Lys				
			8450		8455
30	Ala Ile Asn Asn Ala Asp Thr Val Ser Asp Val Thr Asp Ile Val Asn				
			8465		8470
	Asn Gln Ile Asp Leu Asn Asp Ala Met Glu Thr Leu Lys His Leu Val				
			8485		8490
35	Asp Asn Glu Ile Pro Asn Ala Glu Gln Thr Val Asn Tyr Gln Asn Ala				
			8500		8505
	Asp Asp Asn Ala Lys Thr Asn Phe Asp Asp Ala Lys Arg Leu Ala Asn				
			8515		8520
40	Thr Leu Leu Asn Ser Asp Asn Thr Asn Val Asn Asp Ile Asn Gly Ala				
			8530		8535
	Ile Gln Ala Val Asn Asp Ala Ile His Asn Leu Asn Gly Asp Gln Arg				
			8545		8550
45	Leu Gln Asp Ala Lys Asp Lys Ala Ile Gln Ser Ile Asn Gln Ala Leu				
			8565		8570
	Ala Asn Lys Leu Lys Glu Ile Glu Ala Ser Asn Ala Thr Asp Gln Asp				
			8580		8585
50	Lys Leu Ile Ala Lys Asn Lys Ala Glu Glu Leu Ala Asn Ser Ile Ile				
			8595		8600
	Asn Asn Ile Asn Lys Ala Thr Ser Asn Gln Ala Val Ser Gln Val Gln				
			8610		8615
55	Thr Ala Gly Asn His Ala Ile Glu Gln Val His Ala Asn Glu Ile Pro				
			8625		8630
					8635
					8640

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Lys Ala Lys Ile Asp Ala Asn Lys Asp Val Asp Lys Gln Val Gln Ala
 8645 8650 8655
 5 Leu Ile Asp Glu Ile Asp Arg Asn Pro Asn Leu Thr Asp Lys Glu Lys
 8660 8665 8670
 Gln Ala Leu Lys Asp Arg Ile Asn Gln Ile Leu Gln Gln Gly His Asn
 8675 8680 8685
 10 Gly Ile Asn Asn Ala Met Thr Lys Glu Glu Ile Glu Gln Ala Lys Ala
 8690 8695 8700
 Gln Leu Ala Gln Ala Leu Gln Asp Ile Lys Asp Leu Val Lys Ala Lys
 8705 8710 8715 8720
 15 Glu Asp Ala Lys Gln Asp Val Asp Lys Gln Val Gln Ala Leu Ile Asp
 8725 8730 8735
 Glu Ile Asp Gln Asn Pro Asn Leu Thr Asp Lys Glu Lys Gln Ala Leu
 8740 8745 8750
 20 Lys Asp Arg Ile Asn Gln Ile Leu Gln Gln Gly His Asn Asp Ile Asn
 8755 8760 8765
 Asn Ala Met Thr Lys Glu Ala Ile Glu Gln Ala Lys Glu Arg Leu Ala
 8770 8775 8780
 25 Gln Ala Leu Gln Asp Ile Lys Asp Leu Val Lys Ala Lys Glu Asp Ala
 8785 8790 8795 8800
 Lys Asn Asp Ile Asp Lys Arg Val Gln Ala Leu Ile Asp Glu Ile Asp
 8805 8810 8815
 30 Gln Asn Pro Asn Leu Thr Asp Lys Glu Lys Gln Ala Leu Lys Asp Arg
 8820 8825 8830
 Ile Asn Gln Ile Leu Gln Gln Gly His Asn Asp Ile Asn Asn Ala Leu
 8835 8840 8845
 35 Thr Lys Glu Glu Ile Glu Gln Ala Lys Ala Gln Leu Ala Gln Ala Leu
 8850 8855 8860
 Gln Asp Ile Lys Asp Leu Val Lys Ala Lys Glu Asp Ala Lys Asn Ala
 8865 8870 8875 8880
 40 Ile Lys Ala Leu Ala Asn Ala Lys Arg Asp Gln Ile Asn Ser Asn Pro
 8885 8890 8895
 Asp Leu Thr Pro Glu Gln Lys Ala Lys Ala Leu Lys Glu Ile Asp Glu
 8900 8905 8910
 45 Ala Glu Lys Arg Ala Leu Gln Asn Val Glu Asn Ala Gln Thr Ile Asp
 8915 8920 8925
 Gln Leu Asn Arg Gly Leu Asn Leu Gly Leu Asp Asp Ile Arg Asn Thr
 8930 8935 8940
 50 His Val Trp Glu Val Asp Glu Gln Pro Ala Val Asn Glu Ile Phe Glu
 8945 8950 8955 8960
 Ala Thr Pro Glu Gln Ile Leu Val Asn Gly Glu Leu Ile Val His Arg
 8965 8970 8975
 55 Asp Asp Ile Ile Thr Glu Gln Asp Ile Leu Ala His Ile Asn Leu Ile
 8980 8985 8990
 Asp Gln Leu Ser Ala Glu Val Ile Asp Thr Pro Ser Thr Ala Thr Ile
 8995 9000 9005

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Ser Asp Ser Leu Thr Ala Lys Val Glu Val Thr Leu Leu Asp Gly Ser
 9010 9015 9020
 5 Lys Val Ile Val Asn Val Pro Val Lys Val Val Glu Lys Glu Leu Ser
 9025 9030 9035
 Val Val Lys Gln Gln Ala Ile Glu Ser Ile Glu Asn Ala Ala Gln Gln
 9045 9050 9055
 10 Lys Ile Asn Glu Ile Asn Asn Ser Val Thr Leu Thr Leu Glu Gln Lys
 9060 9065 9070
 Glu Ala Ala Ile Ala Glu Val Asn Lys Leu Lys Gln Gln Ala Ile Asp
 9075 9080 9085
 15 His Val Asn Asn Ala Pro Asp Val His Ser Val Glu Glu Ile Gln Gln
 9090 9095 9100
 Gln Glu Gln Ala His Ile Glu Gln Phe Asn Pro Glu Gln Phe Thr Ile
 9105 9110 9115 9120
 20 Glu Gln Ala Lys Ser Asn Ala Ile Lys Ser Ile Glu Asp Ala Ile Gln
 9125 9130 9135
 His Met Ile Asp Glu Ile Lys Ala Arg Thr Asp Leu Thr Asp Lys Glu
 9140 9145 9150
 25 Lys Gln Glu Ala Ile Ala Lys Leu Asn Gln Leu Lys Glu Gln Ala Ile
 9155 9160 9165
 Gln Ala Ile Gln Arg Ala Gln Ser Ile Asp Glu Ile Ser Glu Gln Leu
 9170 9175 9180
 Glu Gln Phe Lys Ala Gln Met Lys Ala Ala Asn Pro Thr Ala Lys Glu
 9185 9190 9195 9200
 30 Leu Ala Lys Arg Lys Gln Glu Ala Ile Ser Arg Ile Lys Asp Phe Ser
 9205 9210 9215
 Asn Glu Lys Ile Asn Ser Ile Arg Asn Ser Glu Ile Gly Thr Ala Asp
 9220 9225 9230
 35 Glu Lys Gln Ala Ala Met Asn Gln Ile Asn Glu Ile Val Leu Glu Thr
 9235 9240 9245
 Ile Arg Asp Ile Asn Asn Ala His Thr Leu Gln Gln Val Glu Ala Ala
 9250 9255 9260
 40 Leu Asn Asn Gly Ile Ala Arg Ile Ser Ala Val Gln Ile Val Thr Ser
 9265 9270 9275 9280
 Asp Arg Ala Lys Gln Ser Ser Ser Thr Gly Asn Glu Ser Asn Ser His
 9285 9290 9295
 45 Leu Thr Ile Gly Tyr Gly Thr Ala Asn His Pro Phe Asn Ser Ser Thr
 9300 9305 9310
 Ile Gly His Lys Lys Lys Leu Asp Glu Asp Asp Asp Ile Asp Pro Leu
 9315 9320 9325
 50 His Met Arg His Phe Ser Asn Asn Phe Gly Asn Val Ile Lys Asn Ala
 9330 9335 9340
 Ile Gly Val Val Gly Ile Ser Gly Leu Leu Ala Ser Phe Trp Phe Phe
 9345 9350 9355 9360
 55 Ile Ala Lys Arg Arg Arg Lys Glu Asp Glu Glu Glu Glu Leu Glu Ile
 9365 9370 9375
 Arg Asp Asn Asn Lys Asp Ser Ile Lys Glu Thr Leu Asp Asp Thr Lys

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	9380	9385	9390
	His Leu Pro Leu Leu Phe Ala Lys Arg Arg Arg Lys Glu Asp Glu Glu		
	9395	9400	9405
5	Asp Val Thr Val Glu Glu Lys Asp Ser Leu Asn Asn Gly Glu Ser Leu		
	9410	9415	9420
	Asp Lys Val Lys His Thr Pro Phe Phe Leu Pro Lys Arg Arg Arg Lys		
	9425	9430	9435
10	Glu Asp Glu Glu Asp Val Glu Val Thr Asn Glu Asn Thr Asp Glu Lys		
	9445	9450	9455
	Val Leu Lys Asp Asn Glu His Ser Pro Leu Leu Phe Ala Lys Arg Arg		
	9460	9465	9470
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	Pro Pro Arg Asn Ala Gln Arg Arg Lys Arg Arg Arg Asp Leu Ala Thr		
	50 55 60		
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	Val Gln Asn Glu Ala Gly Thr Ile Asp Asp Arg Gln Val Glu Ser Ser		
	85 90 95		
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	Glu Gln Ser Gln Gln Pro Lys Pro Tyr Phe Thr Thr Gly Ala Asn Gln		
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55	Ser Glu Thr Ser Lys Asn Glu His Asp Asn Asp Ser Val Lys Gln Asp		
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5 Gln Asp Glu Pro Lys Glu His His Asn Gly Lys Lys Ala Ala Ala Ile
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 Gly Ala Gly Thr Ala Gly Val Ala Gly Ala Ala Gly Ala Met Ala Ala
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 Ser Lys Ala Lys Lys His Ser Asn Asp Ala Gln Asn Lys Ser Asn Ser
 225 230 235 240
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 Lys Asp His His Asn Gly Lys Lys Gly Ala Ala Ile Gly Ala Gly Thr
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 15 Ala Gly Leu Ala Gly Gly Ala Ala Ser Lys Ser Ala Ser Ala Ala Ser
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 290 295 300
 20 Asn His Asp Arg Asp Lys Glu Arg Lys Lys Gly Gly Met Ala Lys Val
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 Leu Leu Pro Leu Ile Ala Ala Val Leu Ile Ile Gly Ala Leu Ala Ile
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Tyr Gly Pro Arg Glu Lys Lys Pro Val Ser Ile Asn His Asn Ile Val
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 Glu Tyr Asn Asp Gly Thr Phe Lys Tyr Gln Ser Arg Pro Lys Phe Asn
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 Ser Thr Pro Lys Tyr Ile Lys Phe Lys His Asp Tyr Asn Ile Leu Glu
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 Phe Asn Asp Gly Thr Phe Glu Tyr Gly Ala Arg Pro Gln Phe Asn Lys
 10 85 90 95
 Pro Ala Ala Lys Thr Asp Ala Thr Ile Lys Lys Glu Gln Lys Leu Ile
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 Gln Ala Gln Asn Leu Val Arg Glu Phe Glu Lys Thr His Thr Val Ser
 15 115 120 125
 Ala His Arg Lys Ala Gln Lys Ala Val Asn Leu Val Ser Phe Glu Tyr
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 Thr Asp Asn Lys Asn Phe Val Ala Ser Glu Asp Lys Leu Asn Lys Ile
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 Ala Asp Ser Ser Ala Ala Ser Lys Ile Val Asp Lys Asn Phe Val Val
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 Pro Glu Ser Lys Leu Gly Asn Ile Val Pro Glu Tyr Lys Glu Ile Asn
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 180 185 190
 Pro Arg Tyr Thr His Pro Ser Gln Ser Leu Ile Ile Lys His His Phe

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 10 Phe Ala Gln Leu Leu Glu Glu Ile Lys Gln Gln Leu Asn Ser Thr Ala
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 65 70 75 80
 55 Asn Ile Glu Thr Val Lys Glu Glu Val Val Lys Glu Glu Ala Lys Pro
 85 90 95

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5 Gln Val Lys Glu Thr Thr Gln Ser Gln Asp Asn Ser Gly Asp Gln Arg
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 5 Gln Val Asp Leu Thr Pro Lys Lys Ala Thr Gln Asn Gln Val Ala Glu
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 Thr Gln Val Glu Val Ala Gln Pro Arg Thr Ala Ser Gly Ser Lys Pro
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 10 Arg Val Thr Arg Ser Ala Asp Val Ala Glu Ala Lys Glu Ala Ser Asn
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 Ala Lys Val Glu Thr Gly Thr Asp Val Thr Ser Lys Val Thr Val Glu
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 15 Ile Gly Ser Ile Glu Gly His Asn Asn Thr Asn Lys Val Glu Pro His
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 195 200 205
 20 Leu His Gln Gly Asp Tyr Phe Asp Phe Thr Leu Ser Asn Val Asn
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 Thr His Gly Val Ser Thr Ala Arg Lys Val Pro Glu Ile Lys Asn Gly
 225 230 235 240
 Ser Val Val Met Ala Thr Gly Glu Val Leu Glu Gly Gly Lys Ile Arg
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 25 Tyr Thr Phe Thr Asn Asp Ile Glu Asp Lys Val Asp Val Thr Ala Glu
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 Val Arg Ile Phe Glu Tyr Leu Gly Asn Asn Glu Asp Ile Ala Lys Ser
 370 375 380
 45 Val Tyr Ala Asn Thr Thr Asp Thr Ser Lys Phe Lys Glu Val Thr Ser
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 Tyr Leu Asn Gly Thr Asp Glu Val Asp Phe Arg Thr Gln Met Val Gly
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 55 His Pro Glu Gln Leu Tyr Lys Tyr Tyr Tyr Asp Arg Gly Tyr Thr Leu
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40	Gly	Asn	Gln	Ser	Phe	Glu	Glu	Asp	Thr	Glu	Glu	Asp	Lys	Pro	Lys	Tyr
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	Glu	Gln	Gly	Gly	Asn	Ile	Val	Asp	Ile	Asp	Phe	Asp	Ser	Val	Pro	Gln
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		770					775					780				
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 885 890 895
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 945 950 960
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 Glu Thr Thr Gly Thr Gln Val Asp Ile Ala Gln Pro Ser Asn Val Ser
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 Asp Thr Asn Val Val Asn Pro His Asn Ala Glu Arg Val Thr Leu Lys
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 55 Tyr Lys Trp Lys Phe Gly Glu Gly Ile Lys Ala Gly Asp Tyr Phe Asp
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 Glu Ile Ile Gly Glu Arg Lys Val Arg Tyr Thr Phe Lys Glu Tyr Val
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 Asp Pro Thr Thr Val Thr Gln Lys Gly Asn Gln Asn Val Glu Val Lys
 260 265 270
 15 Leu Gly Glu Thr Thr Val Ser Lys Ile Phe Asn Ile Gln Tyr Leu Gly
 275 280 285
 Gly Val Arg Asp Asn Trp Gly Val Thr Ala Asn Gly Arg Ile Asp Thr
 290 295 300
 20 Leu Asn Lys Val Asp Gly Lys Phe Ser His Phe Ala Tyr Met Lys Pro
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 Asn Asn Gln Ser Leu Ser Ser Val Thr Val Thr Gly Gln Val Thr Lys
 325 330 335
 25 Gly Asn Lys Pro Gly Val Asn Asn Pro Thr Val Lys Val Tyr Lys His
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 Val Ser Lys Phe Glu Asp Val Thr Asp Asn Met Ser Leu Asp Phe Asp
 370 375 380
 30 Thr Asn Gly Gly Tyr Ser Leu Asn Phe Asn Asn Leu Asp Gln Ser Lys
 385 390 395 400
 Asn Tyr Val Ile Lys Tyr Glu Gly Tyr Tyr Asp Ser Asn Ala Ser Asn
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 35 Leu Glu Phe Gln Thr His Leu Phe Gly Tyr Tyr Asn Tyr Tyr Tyr Thr
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 Ser Asn Leu Thr Trp Lys Asn Gly Val Ala Phe Tyr Ser Asn Asn Ala
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 15 Phe Tyr Ser Phe Ile Asp Asp Lys Asn His Asn Lys Lys Leu Leu Val
 65 70 75 80
 Ile Arg Thr Lys Gly Thr Ile Ala Gly Gln Tyr Arg Val Tyr Ser Glu
 85 90 95
 20 Glu Gly Ala Asn Lys Ser Gly Leu Ala Trp Pro Ser Ala Phe Lys Val
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 Gln Leu Gln Leu Pro Asp Asn Glu Val Ala Gln Ile Ser Asp Tyr Tyr
 115 120 125
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 35 Trp Lys Val Ile Phe Asn Asn Met Val Asn Gln Asn Trp Gly Pro Tyr
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 210 215 220
 40 Thr Arg Asn Gly Ser Met Lys Ala Ala Asp Asn Phe Leu Asp Pro Asn
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 Lys Ala Ser Ser Leu Leu Ser Ser Gly Phe Ser Pro Asp Phe Ala Thr
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 45 Val Ile Thr Met Asp Arg Lys Ala Ser Lys Gln Gln Thr Asn Ile Asp
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 10 Ala Thr Ala Asp Ser Asp Lys Phe Lys Ile Ser Gln Ile Leu Thr Phe
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 Asn Gln Asn Glu Glu Phe Gln Val Gln Asn Thr Leu Gly Tyr Thr Phe
 Gly Gly Asp Ile Ser Ile Ser Asn Gly Leu Ser Gly Gly Leu Asn Gly
 25 Asn Thr Ala Phe Ser Glu Thr Ile Asn Tyr Lys Gln Glu Ser Tyr Arg
 Thr Thr Leu Ser Arg Asn Thr Asn Tyr Lys Asn Val Gly Trp Gly Val
 30 Glu Ala His Lys Ile Met Asn Asn Gly Trp Gly Pro Tyr Gly Arg Asp
 Ser Phe His Pro Thr Tyr Gly Asn Glu Leu Phe Leu Ala Gly Arg Gln
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 Arg Glu Met Asp Leu Tyr Gln Ile Arg Trp Asn Gly Phe Tyr Trp Ala
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 5 Glu Ala Thr Asn Ala Thr Asn Asn Gln Ser Thr Gln Val Ser Gln Ala
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 10 Lys Ser His Met Asp Asp Tyr Met Gln His Pro Gly Lys Val Ile Lys
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 15 Trp Lys Glu Tyr Lys Phe Tyr Asn Ala Asn Asn Gln Glu Leu Ala Thr
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 145 150 155 160
 Val Pro Gln Ile Asn Tyr Asn His Arg Tyr Thr Thr His Leu Glu Phe
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 25 Glu Lys Ala Ile Pro Thr Leu Ala Asp Ala Ala Lys Pro Asn Asn Val
 180 185 190
 Lys Pro Val Gln Pro Lys Pro Ala Gln Pro Lys Thr Pro Thr Glu Gln
 195 200 205
 30 Thr Lys Pro Val Gln Pro Lys Val Glu Lys Val Lys Pro Thr Val Thr
 210 215 220
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 245 250 255
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 260 265 270
 40 Asp Val Ala Thr Ala Lys Ser Glu Ser Asn Asn Gln Ala Val Ser Asp
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 45 Lys Gln Ala Ser Lys Ala Lys Glu Leu Pro Lys Thr Gly Leu Thr Ser
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 260 265 270
 40 Tyr Lys Ala Glu Lys Leu Leu Ala Pro Tyr Lys Lys Ala Lys Thr Leu
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 45 Lys Leu Lys Ala Glu Tyr Lys Lys Lys Leu Glu Asp Thr Lys Lys Ala
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5 Lys Asp Glu Arg Thr Ser Glu Phe Glu Val Ser Lys Leu Asn Gly Lys
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15 Ser Ala Ser Gly Ser Asp Lys Gly Ser Asp Gly Thr Thr Thr Gly Gln
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40 Phe Asp Glu Val Lys Ile Leu Thr Val Trp Lys Ser Lys Gln Ala Phe
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45 Thr Asp Trp Leu Lys Ser Asp Val Phe Lys Ala Ala His Lys His Val
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 Pro Ala Leu Glu Asn Lys Glu His Asp Ile Gly Pro Arg Glu Gln Val
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 Asn Phe Gln Leu Leu Asp Lys Asn Asn Glu Thr Gln Tyr Tyr His Phe
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 Glu Val Glu Leu Asp Ile Asn Thr Ala Ser Thr Trp Lys Lys Phe Glu
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 Val Tyr Glu Asn Asn Gln Lys Leu Pro Val Arg Leu Val Ser Tyr Ser
 165 170 175
 Pro Val Pro Glu Asp His Ala Tyr Ile Arg Phe Pro Val Ser Asp Gly
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 Thr Gln Glu Leu Lys Ile Val Ser Ser Thr Gln Ile Asp Asp Gly Glu
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 Glu Thr Asn Tyr Asp Tyr Thr Lys Leu Val Phe Ala Lys Pro Ile Tyr
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 Ser Asn Gln Asn Ile Ser Thr Ile Asn Asn Ala Asn Asn Gln Pro Gln
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 Gln Gln Tyr Pro Pro Ala Asp Glu Ser Leu Gln Asp Ala Ile Lys Asn
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 Pro Ala Ile Ile Asp Lys Glu His Thr Ala Asp Asn Trp Arg Pro Ile
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 370 375 380
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 Val Tyr Glu Gly Asp Lys Lys Leu Pro Val Glu Leu Val Ser Tyr Asp

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 5 Gly Lys Ala Ala Lys Leu Asp Val Val Lys Gln Asn Tyr Asn Asn Thr
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 Asp Lys Val Thr Asp Lys Lys Thr Thr Glu His Leu Pro Ser Asp Ile
 820 825 830
 10 His Lys Thr Val Asp Lys Thr Val Lys Thr Lys Glu Lys Ala Gly Thr
 835 840 845
 Pro Ser Lys Glu Asn Lys Leu Ser Gln Ser Lys Met Leu Pro Lys Thr
 850 855 860
 15 Gly Glu Thr Thr Ser Ser Gln Ser Trp Trp Gly Leu Tyr Ala Leu Leu
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 Asp Glu Val Lys Ile Leu Thr Ile Trp Glu Ser Glu Asp Ser Phe Asn
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 35 Asn Trp Leu Asn Ser Asp Val Phe Lys Glu Ala His Lys Asn Val Arg
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Thr Thr Ile Asp Arg Lys Thr Asn His Lys Ser Ile Gly Trp Gly Val
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195 200 205

Ser Tyr Asp Pro Thr Tyr Gly Asn Glu Leu Phe Leu Gly Gly Arg Gln
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Leu Leu Ala Arg Gly Asn Phe Asn Pro Glu Phe Ile Ser Val Leu Ser
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Arg Glu Met Asp Arg Tyr Thr Asn Gln Trp Asn Arg Leu His Trp Val
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35 Gly Asn Asn Tyr Lys Asn Gln Asn Thr Val Thr Phe Thr Ser Thr Tyr
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Val Lys Gly Ser Gly Tyr Glu Leu Thr Lys Arg Met Ile Trp Pro Phe

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Thr Asp Phe Ala Pro Lys Asn Gln Asp Glu Ser Arg Glu Val Lys Tyr
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 5 Thr Tyr Gly Tyr Lys Thr Gly Gly Asp Phe Ser Ile Asn Arg Gly Gly
 145 150 155 160
 Leu Thr Gly Asn Ile Thr Lys Glu Ser Asn Tyr Ser Glu Thr Ile Ser
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 10 Tyr Gln Gln Pro Ser Tyr Arg Thr Leu Leu Asp Gln Ser Thr Ser His
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 15 His Asp His Thr Arg Gln Leu Thr Asn Asp Ser Asp Asn Arg Thr Lys
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 Asn Tyr Ser Lys Lys Thr Leu Thr Asn Gln Asp Val Tyr Leu Glu Tyr

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	Ala	Met	Ser	Thr	Phe 245	Ala	Ser	Ala	Ala	Thr 250	Thr	Thr	Ala	Val	Thr 255	Ala	

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Asn Thr Ile Thr Val Asn Lys Asp Asn Leu Lys Gln Tyr Met Thr Thr
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 5 Ser Gly Asn Ala Thr Tyr Asp Gln Ser Thr Gly Ile Val Thr Leu Thr
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 Gln Asp Ala Tyr Ser Gln Lys Gly Ala Ile Thr Leu Gly Thr Arg Ile
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 10 Asp Ser Asn Lys Ser Phe His Phe Ser Gly Lys Val Asn Leu Gly Asn
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 Lys Tyr Glu Gly His Gly Asn Gly Gly Asp Gly Ile Gly Phe Ala Phe
 325 330 335
 15 Ser Pro Gly Val Leu Gly Glu Thr Gly Leu Asn Gly Ala Ala Val Gly
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 20 Asn Thr Ser Lys Pro Asn Ser Ala Ala Lys Ala Asn Ala Asp Pro Ser
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 25 Tyr Gly Val Ala Thr Thr Tyr Thr Ser Ser Ser Thr Ala Asp Asn Ala
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 Ser Gly Leu Thr Phe Asp Ser Thr Asn Asn Thr Ile Ser Gly Thr Pro
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 Gly Asn Lys Thr Thr Thr Thr Phe Lys Tyr Glu Val Thr Arg Asn Ser
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 Val Ser Thr Ser Lys Ala Asp Ser Gln Ser Ala Ser Thr Ser Thr Ser
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 5 Thr Val Thr Asp Lys Val Asn Gly Tyr Ser Leu Ile Asn Asn Gly Lys
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 5 Glu Asp Ala Asn His Val Lys Thr Ala Asn Arg Ala Ser Gln Ala Asp
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 15 Thr Lys Ile Asn Asn Asp Lys Asn Asn Ala Ile Ala Glu Ile Asn Lys
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 Leu Glu Arg Val Lys Asn Gln Glu Ile Ser Lys Ile Glu Asn Ile Thr
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 45 Asp Ser Thr Gln Thr Lys Met Asp Ala Tyr Asn Glu Val Lys Gln Ala
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 Ala Thr Ala Arg Lys Ala Gln Asn Ala Thr Val Ser Asn Ala Thr Asn
 850 855 860
 50 Glu Glu Val Ala Glu Ala Asp Ala Ala Val Asp Ala Ala Gln Lys Gln
 865 870 875 880
 Gly Leu His Asp Ile Gln Val Val Lys Ser Lys Gln Glu Val Ala Asp
 885 890 895
 55 Thr Lys Ser Lys Val Leu Asp Lys Ile Asn Ala Ile Gln Thr Gln Ala
 900 905 910
 Lys Val Lys Pro Ala Ala Asp Thr Glu Val Glu Asn Ala Tyr Asn Thr
 915 920 925
 Arg Lys Gln Glu Ile Gln Asn Ser Asn Ala Ser Thr Thr Glu Glu Lys

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	930					935								940					
	Gln	Ala	Ala	Tyr	Thr	Glu	Leu	Asp	Thr	Lys	Lys	Gln	Glu	Ala	Arg	Thr			
	945					950					955					960			
5	Asn	Leu	Asp	Ala	Ala	Asn	Thr	Asn	Ser	Asp	Val	Thr	Thr	Ala	Lys	Asp			
				965						970					975				
	Asn	Ser	Ile	Ala	Ala	Ile	Asn	Gln	Val	Gln	Ala	Ala	Thr	Thr	Lys	Lys			
				980					985					990					
10	Ser	Asp	Ala	Lys	Ala	Glu	Ile	Ala	Gln	Lys	Ala	Ser	Glu	Arg	Lys	Thr			
			995					1000					1005						
	Ala	Ile	Glu	Ala	Met	Asn	Asp	Ser	Thr	Thr	Glu	Glu	Gln	Gln	Ala	Ala			
		1010					1015				1020								
15	Lys	Asp	Lys	Val	Asp	Gln	Ala	Val	Val	Thr	Ala	Asn	Ala	Asp	Ile	Asp			
	1025					1030					1035				1040				
	Asn	Ala	Ala	Ala	Asn	Asn	Asp	Val	Asp	Asn	Ala	Lys	Thr	Thr	Asn	Glu			
					1045					1050					1055				
20	Ala	Thr	Ile	Ala	Ala	Ile	Thr	Pro	Asp	Ala	Asn	Val	Lys	Pro	Ala	Ala			
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	Lys	Gln	Ala	Ile	Ala	Asp	Lys	Val	Gln	Ala	Gln	Glu	Thr	Ala	Ile	Asp			
			1075					1080					1085						
25	Gly	Asn	Asn	Gly	Ser	Thr	Thr	Glu	Glu	Lys	Ala	Ala	Ala	Lys	Gln	Gln			
		1090					1095					1100							
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	Glu	Ala	Ile	Gln	Pro	Ala	Thr	Thr	Thr	Lys	Asp	Asn	Ala	Lys	Glu	Ala			
				1140					1145					1150					
35	Ile	Ala	Thr	Lys	Ala	Asn	Glu	Arg	Lys	Thr	Ala	Ile	Ala	Gln	Thr	Gln			
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		1170					1175					1180							
40	Ala	Val	Thr	Gln	Ala	Asn	Ser	Asn	Ile	Glu	Ala	Ala	Asn	Ser	Gln	Asn			
	1185					1190					1195				1200				
	Asp	Val	Asp	Gln	Ala	Lys	Thr	Thr	Gly	Glu	Asn	Ser	Ile	Asp	Gln	Val			
					1205					1210					1215				
45	Thr	Pro	Thr	Val	Asn	Lys	Lys	Ala	Thr	Ala	Arg	Asn	Glu	Ile	Thr	Ala			
				1220					1225					1230					
	Ile	Leu	Asn	Asn	Lys	Leu	Gln	Glu	Ile	Gln	Ala	Thr	Pro	Asp	Ala	Thr			
			1235					1240					1245						
50	Asp	Glu	Glu	Lys	Gln	Ala	Ala	Asp	Ala	Glu	Ala	Asn	Thr	Glu	Asn	Gly			
		1250					1255					1260							
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	1265					1270					1275					1280			
	Glu	Ala	Lys	Ala	Asn	Ala	Glu	Ala	Ala	Ile	Asn	Ala	Val	Thr	Pro	Lys			
					1285					1290					1295				
55	Val	Val	Lys	Lys	Gln	Ala	Ala	Lys	Asp	Glu	Ile	Asp	Gln	Leu	Gln	Ala			
				1300					1305					1310					

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Thr Gln Thr Asn Val Ile Asn Asn Asp Gln Asn Ala Thr Thr Glu Glu
 1315 1320 1325
 Lys Glu Ala Ala Ile Gln Gln Leu Ala Thr Ala Val Thr Asp Ala Lys
 5 1330 1335 1340
 Asn Asn Ile Thr Ala Ala Thr Asp Asp Asn Gly Val Asp Gln Ala Lys
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 Asp Ala Gly Lys Asn Ser Ile Gln Ser Thr Gln Pro Ala Thr Ala Val
 10 1365 1370 1375
 Lys Ser Asn Ala Lys Asn Asp Val Asp Gln Ala Val Thr Thr Gln Asn
 1380 1385 1390
 Gln Ala Ile Asp Asn Thr Thr Gly Ala Thr Thr Glu Glu Lys Asn Ala
 15 1395 1400 1405
 Ala Lys Asp Leu Val Leu Lys Ala Lys Glu Lys Ala Tyr Gln Asp Ile
 1410 1415 1420
 Leu Asn Ala Gln Thr Thr Asn Asp Val Thr Gln Ile Lys Asp Gln Ala
 20 1425 1430 1435 1440
 Val Ala Asp Ile Gln Gly Ile Thr Ala Asp Thr Thr Ile Lys Asp Val
 1445 1450 1455
 Ala Lys Asp Glu Leu Ala Thr Lys Ala Asn Glu Gln Lys Ala Leu Ile
 25 1460 1465 1470
 Ala Gln Thr Ala Asp Ala Thr Thr Glu Glu Lys Glu Gln Ala Asn Gln
 1475 1480 1485
 Gln Val Asp Ala Gln Leu Thr Gln Gly Asn Gln Asn Ile Glu Asn Ala
 30 1490 1495 1500
 Gln Ser Ile Asp Asp Val Asn Thr Ala Lys Asp Asn Ala Ile Gln Ala
 1505 1510 1515 1520
 Ile Asp Pro Ile Gln Ala Ser Thr Asp Val Lys Thr Asn Ala Arg Ala
 35 1525 1530 1535
 Glu Leu Leu Thr Glu Met Gln Asn Lys Ile Thr Glu Ile Leu Asn Asn
 1540 1545 1550
 Asn Glu Thr Thr Asn Glu Glu Lys Gly Asn Asp Ile Gly Pro Val Arg
 40 1555 1560 1565
 Ala Ala Tyr Glu Glu Gly Leu Asn Asn Ile Asn Ala Ala Thr Thr Thr
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 Gly Asp Val Thr Thr Ala Lys Asp Thr Ala Val Gln Lys Val Gln Gln
 45 1585 1590 1595 1600
 Leu His Ala Asn Pro Val Lys Lys Pro Ala Gly Lys Lys Glu Leu Asp
 1605 1610 1615
 Gln Ala Ala Ala Asp Lys Lys Thr Gln Ile Glu Gln Thr Pro Asn Ala
 50 1620 1625 1630
 Ser Gln Gln Glu Ile Asn Asp Ala Lys Gln Glu Val Asp Thr Glu Leu
 1635 1640 1645
 Asn Gln Ala Lys Thr Asn Val Asp Gln Ser Ser Thr Asn Glu Tyr Val
 55 1650 1655 1660
 Asp Asn Ala Val Lys Glu Gly Lys Ala Lys Ile Asn Ala Val Lys Thr
 1665 1670 1675 1680

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Phe Ser Glu Tyr Lys Lys Asp Ala Leu Ala Lys Ile Glu Asp Ala Tyr
 1685 1690 1695
 5 Asn Ala Lys Val Asn Glu Ala Asp Asn Ser Asn Ala Ser Thr Ser Ser
 1700 1705 1710
 Glu Ile Ala Glu Ala Lys Gln Lys Leu Ala Glu Leu Lys Gln Thr Ala
 1715 1720 1725
 10 Asp Gln Asn Val Asn Gln Ala Thr Ser Lys Asp Asp Ile Glu Val Gln
 1730 1735 1740
 Ile His Asn Asp Leu Asp Asn Ile Asn Asp Tyr Thr Ile Pro Thr Gly
 1745 1750 1755
 15 Lys Lys Glu Ser Ala Thr Thr Asp Leu Tyr Ala Tyr Ala Asp Gln Lys
 1765 1770 1775
 Lys Asn Asn Ile Ser Ala Asp Thr Asn Ala Thr Gln Asp Glu Lys Gln
 1780 1785 1790
 20 Gln Ala Ile Lys Gln Val Asp Gln Asn Val Gln Thr Ala Leu Glu Ser
 1795 1800 1805
 Ile Asn Asn Gly Val Asp Asn Gly Asp Val Asp Asp Ala Leu Thr Gln
 1810 1815 1820
 25 Gly Lys Ala Ala Ile Asp Ala Ile Gln Val Asp Ala Thr Val Lys Pro
 1825 1830 1835 1840
 Lys Ala Asn Gln Ala Ile Glu Val Lys Ala Glu Asp Thr Lys Glu Ser
 1845 1850 1855
 30 Ile Asp Gln Ser Asp Gln Leu Thr Ala Glu Glu Lys Thr Glu Ala Leu
 1860 1865 1870
 Ala Met Ile Lys Gln Ile Thr Asp Gln Ala Lys Gln Gly Ile Thr Asp
 1875 1880 1885
 Ala Thr Thr Thr Ala Glu Val Glu Lys Ala Lys Ala Gln Gly Leu Glu
 1890 1895 1900
 35 Ala Phe Asp Asn Ile Gln Ile Asp Ser Thr Glu Lys Gln Lys Ala Ile
 1905 1910 1915 1920
 Glu Glu Leu Glu Thr Ala Leu Asp Gln Ile Glu Ala Gly Val Asn Val
 1925 1930 1935
 40 Asn Ala Asp Ala Thr Thr Glu Glu Lys Glu Ala Phe Thr Asn Ala Leu
 1940 1945 1950
 Glu Asp Ile Leu Ser Lys Ala Thr Glu Asp Ile Ser Asp Gln Thr Thr
 1955 1960 1965
 45 Asn Ala Glu Ile Ala Thr Val Lys Asn Ser Ala Leu Glu Gln Leu Lys
 1970 1975 1980
 Ala Gln Arg Ile Asn Pro Glu Val Lys Lys Asn Ala Leu Glu Ala Ile
 1985 1990 1995 2000
 50 Arg Glu Val Val Asn Lys Gln Ile Glu Ile Ile Lys Asn Ala Asp Ala
 2005 2010 2015
 Asp Ala Ser Ala Lys Glu Ile Ala Arg Thr Asp Leu Gly Arg Tyr Phe
 2020 2025 2030
 55 Asp Arg Phe Ala Asp Lys Leu Asp Lys Thr Gln Thr Asn Ala Glu Val
 2035 2040 2045
 Ala Glu Leu Gln Asn Val Thr Ile Pro Ala Ile Glu Ala Ile Val Pro

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	2050					2055						2060					
	Gln	Asn	Asp	Pro	Asp	Ala	Asn	Asp	Thr	Asn	Asn	Gly	Ile	Asp	Asn	Asn	2080
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	Asp	Ala	Thr	Ala	Asn	Ser	Asn	Ala	Asn	Ala	Thr	Pro	Glu	Asn	Thr	Gly	
					2085					2090					2095		
	Gln	Pro	Asn	Val	Ser	Glu	Thr	Thr	Ala	Asn	Gly	Lys	Ala	Asp	Ala	Ser	
				2100					2105					2110			
10	Pro	Thr	Thr	Pro	Asn	Asn	Ser	Asp	Ala	Ala	Thr	Gly	Glu	Thr	Thr	Ala	
			2115					2120					2125				
	Thr	Ser	Ala	Thr	Asp	Asp	Ala	Asn	Asp	Lys	Pro	Gln	Ala	Asn	Asn	Asn	
		2130					2135					2140					
15	Ser	Ser	Val	Asp	Ala	Ser	Thr	Asn	Ser	Pro	Thr	Met	Asp	Asn	Asp	Val	
	2145					2150					2155					2160	
	Thr	Ser	Lys	Pro	Glu	Val	Glu	Ser	Thr	Asn	Asn	Gly	Thr	Thr	Asp	Lys	
					2165										2175		
20	Pro	Val	Thr	Glu	Thr	Asp	Asn	Ala	Thr	Pro	Ala	Glu	Ser	Thr	Thr	Asn	
				2180					2185						2190		
	Asn	Asn	Ser	Thr	Thr	Thr	Ala	Thr	Asn	Glu	Asn	Ala	Pro	Thr	Gly	Ser	
			2195					2200					2205				
25	Thr	Ala	Thr	Ala	Pro	Thr	Thr	Ala	Ser	Thr	Glu	Ala	Ala	Ser	Ser	Ala	
		2210					2215					2220					
	Asp	Ser	Lys	Asp	Asn	Ala	Ser	Val	Asn	Asp	Ser	Lys	Gln	Asn	Ala	Glu	
	2225				2230						2235					2240	
30	Val	Asn	Asn	Ser	Ala	Glu	Ser	Gln	Ser	Thr	Asn	Asp	Lys	Val	Ala	Gln	
					2245						2250				2255		
	Pro	Lys	Ser	Glu	Asn	Lys	Ala	Lys	Ala	Glu	Lys	Asp	Gly	Ser	Asp	Ser	
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35	Thr	Asn	Gln	Ser	Met	Val	Glu	Ser	Thr	Thr	Glu	Thr	Leu	Pro	Ser	Ala	
			2275					2280					2285				
	Asp	Ile	Thr	Glu	Pro	Asn	Val	Pro	Ser	Asn	Thr	Ser	Lys	Asp	Lys	Glu	
		2290					2295					2300					
40	Glu	Ser	Thr	Thr	Asn	Gln	Thr	Asp	Ala	Gly	Gln	Leu	Lys	Ser	Glu	Thr	
	2305					2310					2315					2320	
	Asn	Val	Ala	Ser	Asn	Glu	Ala	Asp	Lys	Ser	Pro	Ser	Lys	Ala	Asp	Thr	
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45	Glu	Val	Ser	Asn	Lys	Pro	Ser	Thr	Ser	Ala	Ser	Ser	Glu	Ala	Lys	Glu	
				2340					2345					2350			
	Lys	Met	Thr	Ser	Thr	Asn	Val	Ser	Gln	Lys	Asp	Asp	Thr	Ala	Thr	Ala	
			2355					2360					2365				
50	Asp	Thr	Asn	Asp	Thr	Gln	Lys	Ser	Val	Gly	Ser	Ala	Ala	Asn	Asn	Lys	
		2370					2375					2380					
	Ala	Thr	Gln	Asn	Asp	Gly	Ala	Asn	Ala	Ser	Pro	Ala	Thr	Val	Ser	Asn	
	2385					2390					2395					2400	
	Gly	Ser	Asn	Ser	Ala	Asn	Gln	Asp	Met	Leu	Asn	Val	Thr	Asn	Thr	Asp	
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55	Asp	His	Gln	Ala	Lys	Thr	Lys	Ser	Ala	Gln	Gln	Gly	Lys	Val	Asn	Lys	
				2420					2425					2430			

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Ala Lys Gln Gln Ala Lys Thr Leu Pro Asp Thr Gly Met Ser His Asn
2435 2440 2445

5 Asp Asp Leu Pro Tyr Ala Glu Leu Ala Leu Gly Ala Gly Met Ala Phe
2450 2455 2460

Leu Ile Arg Arg Phe Thr Lys Lys Asp Gln Gln Thr Glu Glu
2465 2470 2475

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<212> PRT
<213> Staphylococcus aureus

15 <400> 30
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1 5 10 15

Gly Ile Phe Ser Thr Leu Ile Gly Thr Val Leu Leu Leu Ser Asn Pro
20 20 25 30

Asn Gly Ala Gln Ala Leu Thr Thr Asp Asn Asn Val Gln Ser Asp Thr
35 40 45

25 Asn Gln Ala Thr Pro Val Asn Ser Gln Asp Lys Asp Val Ala Asn Asn
50 55 60

Arg Gly Leu Ala Asn Ser Ala Gln Asn Thr Pro Asn Gln Ser Ala Thr
65 70 75 80

Thr Asn Gln Ala Thr Asn Gln Ala Leu Val Asn His Asn Asn Gly Ser
85 90 95

30 Ile Val Asn Gln Ala Thr Pro Thr Ser Val Gln Ser Ser Thr Pro Ser
100 105 110

Ala Gln Asn Asn Asn His Thr Asp Gly Asn Thr Thr Ala Thr Glu Thr
115 120 125

35 Val Ser Asn Ala Asn Asn Asn Asp Val Val Ser Asn Asn Thr Ala Leu
130 135 140

Asn Val Pro Thr Lys Thr Asn Glu Asn Gly Ser Gly Gly His Leu Thr
145 150 155 160

40 Leu Lys Glu Ile Gln Glu Asp Val Arg His Ser Ser Asn Lys Pro Glu
165 170 175

Leu Val Ala Ile Ala Glu Pro Ala Ser Asn Arg Pro Lys Lys Arg Ser
180 185 190

45 Arg Arg Ala Ala Pro Ala Asp Pro Asn Ala Thr Pro Ala Asp Pro Ala
195 200 205

Ala Ala Ala Val Gly Asn Gly Gly Ala Pro Val Ala Ile Thr Ala Pro
210 215 220

50 Tyr Thr Pro Thr Thr Asp Pro Asn Ala Asn Asn Ala Gly Gln Asn Ala
225 230 235 240

Pro Asn Glu Val Leu Ser Phe Asp Asp Asn Gly Ile Arg Pro Ser Thr
245 250 255

55 Asn Arg Ser Val Pro Thr Val Asn Val Val Asn Asn Leu Pro Gly Phe
260 265 270

Thr Leu Ile Asn Gly Gly Lys Val Gly Val Phe Ser His Ala Met Val
275 280 285

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Arg Thr Ser Met Phe Asp Ser Gly Asp Asn Lys Asn Tyr Gln Ala Gln
 290 295 300
 5 Gly Asn Val Ile Ala Leu Gly Arg Ile His Gly Thr Asp Thr Asn Asp
 305 310 315 320
 His Gly Asp Phe Asn Gly Ile Glu Lys Ala Leu Thr Val Asn Pro Asn
 325 330 335
 10 Ser Glu Leu Ile Phe Glu Phe Asn Thr Met Thr Thr Lys Asn Gly Gln
 340 345 350
 Gly Ala Thr Asn Val Ile Ile Lys Asn Ala Asp Thr Asn Asp Thr Ile
 355 360 365
 15 Ala Glu Lys Thr Val Glu Gly Gly Pro Thr Leu Arg Leu Phe Lys Val
 370 375 380
 Pro Asp Asn Val Arg Asn Leu Lys Ile Gln Phe Val Pro Lys Asn Asp
 385 390 395 400
 20 Ala Ile Thr Asp Ala Arg Gly Ile Tyr Gln Leu Lys Asp Gly Tyr Lys
 405 410 415
 Tyr Tyr Ser Phe Val Asp Ser Ile Gly Leu His Ser Gly Ser His Val
 420 425 430
 Phe Val Glu Arg Arg Thr Met Asp Pro Thr Ala Thr Asn Asn Lys Glu
 435 440 445
 25 Phe Thr Val Thr Thr Ser Leu Lys Asn Asn Gly Asn Ser Gly Ala Ser
 450 455 460
 Leu Asp Thr Asn Asp Phe Val Tyr Gln Val Gln Leu Pro Glu Gly Val
 465 470 475 480
 30 Glu Tyr Val Asn Asn Ser Leu Thr Lys Asp Phe Pro Ser Asn Asn Ser
 485 490 495
 Gly Val Asp Val Asn Asp Met Asn Val Thr Tyr Asp Ala Ala Asn Arg
 500 505
 35 Val Ile Thr Ile Lys Ser Thr Gly Gly Gly Thr Ala Asn Ser Pro Ala
 515 520 525
 Arg Leu Met Pro Asp Lys Ile Leu Asp Leu Arg Tyr Lys Leu Arg Val
 530 535 540
 40 Asn Asn Val Pro Thr Pro Arg Thr Val Thr Phe Asn Glu Thr Leu Thr
 545 550 555 560
 Tyr Lys Thr Tyr Thr Gln Asp Phe Ile Asn Ser Ala Ala Glu Ser His
 565 570 575
 45 Thr Val Ser Thr Asn Pro Tyr Thr Ile Asp Ile Ile Met Asn Lys Asp
 580 585 590
 Ala Leu Gln Ala Glu Val Asp Arg Arg Ile Gln Gln Ala Asp Tyr Thr
 595 600 605
 50 Phe Ala Ser Leu Asp Ile Phe Asn Gly Leu Lys Arg Arg Ala Gln Thr
 610 615 620
 Ile Leu Asp Glu Asn Arg Asn Asn Val Pro Leu Asn Lys Arg Val Ser
 625 630 635 640
 55 Gln Ala Tyr Ile Asp Ser Leu Thr Asn Gln Met Gln His Thr Leu Ile
 645 650 655
 Arg Ser Val Asp Ala Glu Asn Ala Val Asn Lys Lys Val Asp Gln Met

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	660					665					670					
	Glu	Asp	Leu	Val	Asn	Gln	Asn	Asp	Glu	Leu	Thr	Asp	Glu	Glu	Lys	Gln
			675					680					685			
5	Ala	Ala	Ile	Gln	Val	Ile	Glu	Glu	His	Lys	Asn	Glu	Ile	Ile	Gly	Asn
		690					695					700				
	Ile	Gly	Asp	Gln	Thr	Thr	Asp	Asp	Gly	Val	Thr	Arg	Ile	Lys	Asp	Gln
	705					710					715					720
10	Gly	Ile	Gln	Thr	Leu	Ser	Gly	Asp	Thr	Ala	Thr	Pro	Val	Val	Lys	Pro
					725					730					735	
	Asn	Ala	Lys	Lys	Ala	Ile	Arg	Asp	Lys	Ala	Thr	Lys	Gln	Arg	Glu	Ile
				740					745					750		
15	Ile	Asn	Ala	Thr	Pro	Asp	Ala	Thr	Glu	Asp	Glu	Ile	Gln	Asp	Ala	Leu
			755					760					765			
	Asn	Gln	Leu	Ala	Thr	Asp	Glu	Thr	Asp	Ala	Ile	Asp	Asn	Val	Thr	Asn
							775					780				
20	Ala	Thr	Thr	Asn	Ala	Asp	Val	Glu	Thr	Ala	Lys	Asn	Asn	Gly	Ile	Asn
					790						795					800
	Thr	Ile	Gly	Ala	Val	Val	Pro	Gln	Val	Thr	His	Lys	Lys	Ala	Ala	Arg
					805					810					815	
25	Asp	Ala	Ile	Asn	Gln	Ala	Thr	Ala	Thr	Lys	Arg	Gln	Gln	Ile	Asn	Ser
				820				825						830		
	Asn	Arg	Glu	Ala	Thr	Gln	Glu	Glu	Lys	Asn	Ala	Ala	Leu	Asn	Glu	Leu
			835					840					845			
30	Thr	Gln	Ala	Thr	Asn	His	Ala	Leu	Glu	Gln	Ile	Asn	Gln	Ala	Thr	Thr
							855					860				
	Asn	Ala	Asn	Val	Asp	Asn	Ala	Lys	Gly	Asp	Gly	Leu	Asn	Ala	Ile	Asn
							870				875					880
35	Pro	Ile	Ala	Pro	Val	Thr	Val	Val	Lys	Gln	Ala	Ala	Arg	Asp	Ala	Val
					885					890					895	
	Ser	His	Asp	Ala	Gln	Gln	His	Ile	Ala	Glu	Ile	Asn	Ala	Asn	Pro	Asp
				900					905					910		
40	Ala	Thr	Gln	Glu	Glu	Arg	Gln	Ala	Ala	Ile	Asp	Lys	Val	Asn	Ala	Ala
								920					925			
	Val	Thr	Ala	Ala	Asn	Thr	Asn	Ile	Leu	Asn	Ala	Asn	Thr	Asn	Ala	Asp
							935					940				
45	Val	Glu	Gln	Val	Lys	Thr	Asn	Ala	Ile	Gln	Gly	Ile	Gln	Ala	Ile	Thr
						950					955					960
	Pro	Ala	Thr	Lys	Val	Lys	Thr	Asp	Ala	Lys	Asn	Ala	Ile	Asp	Lys	Ser
					965					970					975	
50	Ala	Glu	Thr	Gln	His	Asn	Thr	Ile	Phe	Asn	Asn	Asn	Asp	Ala	Thr	Leu
					980				985					990		
	Glu	Glu	Gln	Gln	Ala	Ala	Gln	Gln	Leu	Leu	Asp	Gln	Ala	Val	Ala	Thr
								1000					1005			
55	Ala	Lys	Gln	Asn	Ile	Asn	Ala	Ala	Asp	Thr	Asn	Gln	Glu	Val	Ala	Gln
							1015					1020				
	Ala	Lys	Asp	Gln	Gly	Thr	Gln	Asn	Ile	Val	Val	Ile	Gln	Pro	Ala	Thr
						1030					1035					1040

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Gln Val Lys Thr Asp Thr Arg Asn Val Val Asn Asp Lys Ala Arg Glu
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 5 Ala Ile Thr Asn Ile Asn Ala Thr Thr Gly Ala Thr Arg Glu Glu Lys
 1060 1065 1070
 Gln Glu Ala Ile Asn Arg Val Asn Thr Leu Lys Asn Arg Ala Leu Thr
 1075 1080 1085
 10 Asp Ile Gly Val Thr Ser Thr Thr Ala Met Val Asn Ser Ile Arg Asp
 1090 1095 1100
 Asp Ala Val Asn Gln Ile Gly Ala Val Gln Pro His Val Thr Lys Lys
 1105 1110 1115 1120
 15 Gln Thr Ala Thr Gly Val Leu Asn Asp Leu Ala Thr Ala Lys Lys Gln
 1125 1130 1135
 Glu Ile Asn Gln Asn Thr Asn Ala Thr Thr Glu Glu Lys Gln Val Ala
 1140 1145 1150
 20 Leu Asn Gln Val Asp Gln Glu Leu Ala Thr Ala Ile Asn Asn Ile Asn
 1155 1160 1165
 Gln Ala Asp Thr Asn Ala Glu Val Asp Gln Ala Gln Gln Leu Gly Thr
 1170 1175 1180
 25 Lys Ala Ile Asn Ala Ile Gln Pro Asn Ile Val Lys Lys Pro Ala Ala
 1185 1190 1195 1200
 Leu Ala Gln Ile Asn Gln His Tyr Asn Ala Lys Leu Ala Glu Ile Asn
 1205 1210 1215
 30 Ala Thr Pro Asp Ala Thr Asn Asp Glu Lys Asn Ala Ala Ile Asn Thr
 1220 1225 1230
 Leu Asn Gln Asp Arg Gln Gln Ala Ile Glu Ser Ile Lys Gln Ala Asn
 1235 1240 1245
 35 Thr Asn Ala Glu Val Asp Gln Ala Ala Thr Val Ala Glu Asn Asn Ile
 1250 1255 1260
 Asp Ala Val Gln Val Asp Val Val Lys Lys Gln Ala Ala Arg Asp Lys
 1265 1270 1275 1280
 40 Ile Thr Ala Glu Val Ala Lys Arg Ile Glu Ala Val Lys Gln Thr Pro
 1285 1290 1295
 Asn Ala Thr Asp Glu Glu Lys Gln Ala Ala Val Asn Gln Ile Asn Gln
 1300 1305 1310
 45 Leu Lys Asp Gln Ala Ile Asn Gln Ile Asn Gln Asn Gln Thr Asn Asp
 1315 1320 1325
 Gln Val Asp Thr Thr Thr Asn Gln Ala Val Asn Ala Ile Asp Asn Val
 1330 1335 1340
 50 Glu Ala Glu Val Val Ile Lys Thr Lys Ala Ile Ala Asp Ile Glu Lys
 1345 1350 1355 1360
 Ala Val Lys Glu Lys Gln Gln Gln Ile Asp Asn Ser Leu Asp Ser Thr
 1365 1370 1375
 55 Asp Asn Glu Lys Glu Val Ala Ser Gln Ala Leu Ala Lys Glu Lys Glu
 1380 1385 1390
 Lys Ala Leu Ala Ala Ile Asp Gln Ala Gln Thr Asn Ser Gln Val Asn
 1395 1400 1405

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Gln Ala Ala Thr Asn Gly Val Ser Ala Ile Lys Ile Ile Gln Pro Glu
 1410 1415 1420
 Thr Lys Val Lys Pro Ala Ala Arg Glu Lys Ile Asn Gln Lys Ala Asn
 1425 1430 1435 1440
 5
 Glu Leu Arg Ala Lys Ile Asn Gln Asp Lys Glu Ala Thr Ala Glu Glu
 1445 1450 1455
 Arg Gln Val Ala Leu Asp Lys Ile Asn Glu Phe Val Asn Gln Ala Met
 1460 1465 1470
 10
 Thr Asp Ile Thr Asn Asn Arg Thr Asn Gln Gln Val Asp Asp Thr Thr
 1475 1480 1485
 Ser Gln Ala Leu Asp Ser Ile Ala Leu Val Thr Pro Asp His Ile Val
 1490 1495 1500
 15
 Arg Ala Ala Ala Arg Asp Ala Val Lys Gln Gln Tyr Glu Ala Lys Lys
 1505 1510 1515 1520
 Arg Glu Ile Glu Gln Ala Glu His Ala Thr Asp Glu Glu Lys Gln Val
 1525 1530 1535
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 Ala Leu Asn Gln Leu Ala Asn Asn Glu Lys Arg Ala Leu Gln Asn Ile
 1540 1545 1550
 Asp Gln Ala Ile Ala Asn Asn Asp Val Lys Arg Val Glu Thr Asn Gly
 1555 1560 1565
 25
 Ile Ala Thr Leu Lys Gly Val Gln Pro His Ile Val Ile Lys Pro Glu
 1570 1575 1580
 Ala Gln Gln Ala Ile Lys Ala Ser Ala Glu Asn Gln Val Glu Ser Ile
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 Thr Glu Glu Glu Arg Leu Ala Ala Lys His Leu Val Glu Gln Ala Leu
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	1890						1895					1900				
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	Ser	Ile	Leu	Ala	Gln	Leu	Gln	Asn	Ile	Tyr	Asp	Thr	Ala	Ile	Gly	Gln
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	Tyr	Val	Ala	Asp	Gly	Asn	Arg	Met	Ile	Asp	Glu	Asp	Ala	Thr	Leu	Asn
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Asn Val Ile Ile Thr Thr Asn Val Gly Val Phe Lys Pro His Ala Val
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Tyr Gln Thr Asn Val Asp Ala Asn Gly Val Asn His Gly Gly Ser Glu
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Gln Asn Thr His Glu Pro Thr Ala Thr His Ala Thr Arg Ser Tyr Ala
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Tyr Asp Ala Gln Ala Ala Ser Glu Lys Asp Thr Glu Ile Thr Lys Glu
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Ile Leu Ser Lys Gln Asp Leu Leu Asp Lys Val Asp Lys Ala Ile Arg
50 55 60

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Gln Ile Glu Gln Leu Lys Gln Leu Ser Ala Ser Ser Lys Glu His Tyr
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 Thr Glu Met Asn Gly Gln Ser Asp Ile Asp Ser Lys Leu Asp Gln Leu
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 Lys Lys His Phe Ala Ser Thr Gly Asp Thr Ser Ser Asp Asp Ile Leu

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 Lys Thr Glu Phe Trp Ala Thr Ser Ser Asp Val Leu Lys Leu Lys Ala
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 Lys Tyr Gly Gln Tyr Phe Arg Pro Gly Ser Val Arg Leu Pro Ser Gln
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 Ser Ser Tyr Pro Asn Asn Ile Gly Gln Ile Asn Lys Asp Val Thr Asp
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	Thr 1105	Ile	Thr	Asp	His 1110	Asp	Asp	Phe	Thr	Leu	Asp 1115	Asn	Gly	Tyr	Phe	Glu 1120
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Glu Ala Lys Ala Ala Glu Asn Thr Ser Thr Glu Asn Ala Lys Gln Asp
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Val Lys Lys Gly Asp Thr Met Thr Ile Asn Tyr Asp Lys Asn Val Ile
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 Asp Gly Glu Lys Asp Ser Asn Gly Ser Ser Val Thr Val Lys Ile Asn

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 Asp His His Asn Lys Ala Lys Ala Leu Pro Glu Thr Gly Ser Glu Asn
 10 1125 1130 1135
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 Tyr Glu Ile Leu His Leu Pro Asn Leu Asn Glu Glu Gln Arg Asn Gly
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 Ala Glu Ala Lys Lys Leu Asn Asp Ala Gln Ala Pro Lys Ala Asp Asn
 5 260 265 270
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 Pro Asn Leu Thr Glu Glu Gln Arg Asn Gly Phe Ile Gln Ser Leu Lys
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 Asp Asp Pro Ser Val Ser Lys Glu Ile Leu Ala Glu Ala Lys Lys Leu
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 Asn Asp Ala Gln Ala Pro Lys Glu Glu Asp Asn Asn Lys Pro Gly Lys
 15 325 330 335
 Glu Asp Asn Asn Lys Pro Gly Lys Glu Asp Asn Asn Lys Pro Gly Lys
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 Val Asp Lys Lys Gln Pro Ala Asn His Ala Asp Ala Asn Lys Ala Gln
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	Leu	Ile	Asn	Val	Lys	Asp	Val	Glu	Asn	Val	Thr	Pro	Asn	Lys	Ala	Leu
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 Val Thr Asn Gly Gly Gly Ile Arg Ala Ser Ile Ala Lys Gly Lys Val
 485 490 495
 Thr Arg Tyr Asp Leu Ile Ser Val Leu Pro Phe Gly Asn Thr Ile Ala
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 Gln Arg Met Leu Leu Gly Lys Pro Ala Val Ser Glu Gln Pro Ala Lys
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 Lys Gln Leu Pro Lys Thr Gly Thr Asn Gln Ser Ser Ser Pro Glu Ala
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15 Thr Tyr Lys 65 Gly Gln 70 Lys Leu Lys Gly Ile 75 Ser Phe Glu Asn Ser Asn 80

Gly Glu Trp Ala Tyr 85 Lys Val Thr Gln Gln 90 Lys Ser Gly Glu Glu Ser 95

20 Glu Val Leu Val 100 Ala Asp Lys Asn Lys 105 Lys Val Ile Asn Lys 110 Lys Thr

Glu Lys Glu 115 Asp Thr Met Asn Glu 120 Asn Asp Asn Phe Lys 125 Tyr Ser Asp

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Gly Asp Ile Lys Glu Trp 150 Ser Leu Glu Lys Asp 155 Asp Gly Lys Leu Val 160

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Asn Glu Thr Ser Lys Val 70 Pro Ala Asn Phe Val 75 Lys Leu Asn Asp Ile 80

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Ile Leu Leu Thr 100 Ile Asp Lys Lys Asp 105 Val Ser Ser Val Glu 110 Asp Ser

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	Lys	Glu	Asp	Leu 340	His	Val	Asp	Glu	Ile 345	Tyr	Gly	Ser	Leu	Tyr 350	His	Thr	
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	Gly 370	Lys	Thr	Lys	Phe	Ala	Asn 375	Ala	Val	Val	Lys	Val 380	Asp	Ser	Glu	Leu	
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	Phe	Ser	Phe	Asp	Val 405	Asp	His	Ala	Gly	Phe 410	Arg	Leu	Gln	Asn	Gly 415	Glu	
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	Gly	Asn	Phe 435	Val	Ser	Lys	Asn	Ile 440	Asp	Ile	Tyr	Glu	Ser 445	Pro	Glu	Glu	
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	His 465	Lys	Leu	His	Gly	Asp 470	Lys	Ile	Val	Gly	Tyr 475	Asp	Thr	Asn	Gly	Phe 480	
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 <213> Staphylococcus aureus

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 Lys Gly Ser Lys Asp Thr Val Lys Ile Glu Asn Asn Tyr Lys Met Arg
 35 40 45

20
 Gly Glu Lys Lys Asp Gly Ser Asp Ala Lys Lys Val Lys Glu Thr Val
 50 55 60

25
 Glu Val Pro Lys Asn Pro Lys Asn Ala Val Val Leu Asp Tyr Gly Ala
 65 70 75 80

30
 Leu Asp Val Met Lys Glu Met Gly Leu Ser Asp Lys Val Lys Ala Leu
 85 90 95

35
 Pro Lys Gly Glu Gly Gly Lys Ser Leu Pro Asn Phe Leu Glu Ser Phe
 100 105 110

40
 Lys Asp Asp Lys Tyr Thr Asn Val Gly Asn Leu Lys Glu Val Asn Phe
 115 120 125

45
 Asp Lys Ile Ala Ala Thr Lys Pro Glu Val Ile Phe Ile Ser Gly Arg
 130 135 140

50
 Thr Ala Asn Gln Lys Asn Leu Asp Glu Phe Lys Lys Ala Ala Pro Lys
 145 150 155 160

55
 Ala Lys Ile Val Tyr Val Gly Ala Asp Glu Lys Asn Leu Ile Gly Ser
 165 170 175

60
 Met Lys Gln Asn Thr Glu Asn Ile Gly Lys Ile Tyr Asp Lys Glu Asp
 180 185 190

65
 Lys Ala Lys Glu Leu Asn Lys Asp Leu Asp Asn Lys Ile Ala Ser Met
 195 200 205

70
 Lys Asp Lys Thr Lys Asn Phe Asn Lys Thr Val Met Tyr Leu Leu Val
 210 215 220

75
 Asn Glu Gly Glu Leu Ser Thr Phe Gly Pro Lys Gly Arg Phe Gly Gly
 225 230 235 240

80
 Leu Val Tyr Asp Thr Leu Gly Phe Asn Ala Val Asp Lys Lys Val Ser
 245 250 255

85
 Asn Ser Asn His Gly Gln Asn Val Ser Asn Glu Tyr Val Asn Lys Glu
 260 265 270

90
 Asn Pro Asp Val Ile Leu Ala Met Asp Arg Gly Gln Ala Ile Ser Gly
 275 280 285

95
 Lys Ser Thr Ala Lys Gln Ala Leu Asn Asn Pro Val Leu Lys Asn Val
 290 295 300

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 Lys Ala Ile Lys Glu Asp Lys Val Tyr Asn Leu Asp Pro Lys Leu Trp
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 Tyr Phe Ala Ala Gly Ser Thr Thr Thr Thr Ile Lys Gln Ile Glu Glu
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<213> Staphylococcus aureus

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35 40 45

Asn Glu Leu Ile Lys Tyr Tyr Thr Gln Pro His Phe Ser Leu Ser Gly
50 55 60

Lys Trp Leu Trp Gln Lys Pro Asn Gly Ser Ile His Ala Thr Leu Gln
65 70 75 80

Thr Trp Val Trp Tyr Ser His Ile Gln Val Phe Gly Ser Glu Ser Trp
85 90 95

Gly Asn Ile Asn Gln Leu Arg Asn Lys Tyr Val Asp Ile Phe Gly Thr
100 105 110

Lys Asp Glu Asp Thr Val Glu Gly Tyr Trp Thr Tyr Asp Glu Thr Phe
115 120 125

Thr Gly Gly Val Thr Pro Ala Ala Thr Ser Ser Asp Lys Pro Tyr Arg
130 135 140

Leu Phe Leu Lys Tyr Ser Asp Lys Gln Gln Thr Ile Ile Gly Gly His
145 150 155 160

Glu Phe Tyr Lys Gly Asn Lys Pro Val Leu Thr Leu Lys Glu Leu Asp
165 170 175

Phe Arg Ile Arg Gln Thr Leu Ile Lys Asn Lys Lys Leu Tyr Asn Gly
180 185 190

Glu Phe Asn Lys Gly Gln Ile Lys Ile Thr Ala Asp Gly Asn Asn Tyr
195 200 205

Thr Ile Asp Leu Ser Lys Lys Leu Lys Leu Thr Asp Thr Asn Arg Tyr
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Val Lys Asn Pro Arg Asn Ala Glu Ile Glu Val Ile Leu Glu Lys Ser
225 230 235 240

Asn

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<212> PRT
<213> Staphylococcus aureus

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Tyr Lys Met Asp Asp Gly Lys Thr Val Asp Ile Pro Lys Asp Pro Lys

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		35					40					45				
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	Leu	Lys	Asp	Lys	Phe	Lys	Gly	Val	Thr	Lys	Ile	Gly	Asp	Gly	Asp	Val
					85					90					95	
10	Glu	Lys	Val	Ala	Lys	Glu	Lys	Pro	Asp	Leu	Ile	Ile	Val	Tyr	Ser	Thr
				100					105					110		
	Asp	Lys	Asp	Ile	Lys	Lys	Tyr	Gln	Lys	Val	Ala	Pro	Thr	Val	Val	Val
			115					120					125			
15	Asp	Tyr	Asn	Lys	His	Lys	Tyr	Leu	Glu	Gln	Gln	Glu	Met	Leu	Gly	Lys
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	145					150					155					160
20	Glu	Thr	Thr	Ala	Lys	Asp	Gly	Lys	Glu	Ile	Lys	Lys	Ala	Ile	Gly	Gln
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	Asp	Ala	Thr	Val	Ser	Leu	Phe	Asp	Glu	Phe	Asp	Lys	Lys	Leu	Tyr	Thr
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25	Tyr	Gly	Asp	Asn	Trp	Gly	Arg	Gly	Gly	Glu	Val	Leu	Tyr	Gln	Ala	Phe
			195					200					205			
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	225					230					235					240
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				260					265					270		
	Val	Asp	Ala	Gly	Thr	Tyr	Trp	Tyr	Asn	Asp	Pro	Tyr	Thr	Leu	Asp	Phe
			275					280					285			
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			35					40					45			
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		50					55					60				
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	65					70					75					80

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Asn Gly Tyr Asn Asn Tyr Asn Asn Tyr Asn Asn Gly Tyr Ser Tyr Asn
 85 90 95
 5 Asn Tyr Ser Arg Tyr Asn Asn Tyr Ser Asn Asn Asn Gln Ser Tyr Asn
 100 105 110
 Tyr Asn Asn Tyr Asn Ser Tyr Asn Thr Asn Ser Tyr Arg Thr Gly Gly
 115 120 125
 10 Leu Gly Ala Ser Tyr Ser Thr Ser Ser Asn Asn Val Gln Val Thr Thr
 130 135 140
 Thr Met Ala Pro Ser Ser Asn Gly Arg Ser Ile Ser Ser Gly Tyr Thr
 145 150 155 160
 15 Ser Gly Arg Asn Leu Tyr Thr Ser Gly Gln Cys Thr Tyr Tyr Val Phe
 165 170 175
 Asp Arg Val Gly Gly Lys Ile Gly Ser Thr Trp Gly Asn Ala Ser Asn
 180 185 190
 20 Trp Ala Asn Ala Ala Ala Arg Ala Gly Tyr Thr Val Asn Asn Thr Pro
 195 200 205
 Lys Ala Gly Ala Ile Met Gln Thr Thr Gln Gly Ala Tyr Gly His Val
 210 215 220
 25 Ala Tyr Val Glu Ser Val Asn Ser Asn Gly Ser Val Arg Val Ser Glu
 225 230 235 240
 Met Asn Tyr Gly Tyr Gly Pro Gly Val Val Thr Ser Arg Thr Ile Ser
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 30 Ala Ser Gln Ala Ala Gly Tyr Asn Phe Ile His
 260 265
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 Asp Asp Lys Ala Lys Thr Glu Gln Gln Asp Lys Lys Thr Asn Asp Ser
 35 40 45
 45 Ser Lys Asp Lys Lys Asp Asn Lys Asp Asp Ser Lys Asp Val Asn Lys
 50 55 60
 Asp Asn Lys Asp Asn Ser Ala Asn Asp Asn Gln Gln Gln Ser Asn Ser
 65 70 75 80
 50 Asn Ala Thr Asn Asn Asp Gln Asn Gln Thr Asn Asn Asn Gln Ser Ser
 85 90 95
 55 Asn Asn Gln Ala Asn Asn Asn Gln Lys Ser Ser Tyr Val Ala Pro Tyr
 100 105 110
 Tyr Gly Gln Asn Ala Ala Pro Val Ala Arg Gln Ile Tyr Pro Phe Asn
 115 120 125
 Gly Asn Lys Asn Gln Ala Leu Gln Gln Leu Pro Asn Phe Gln Thr Ala
 130 135 140

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Leu Asn Ala Ala Asn Asn Glu Ala Asn Lys Phe Gly Ser Asn Asn Lys
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 Val Tyr Asn Asp Tyr Ser Ile Glu Glu His Asn Gly Asn Tyr Lys Tyr
 5 165 170 175
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 <213> Staphylococcus aureus
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 Thr Gln Gln Thr Ser Thr Lys His Gln Thr Thr Gln Asn Asn Tyr Val
 35 40 45
 Thr Asp Gln Gln Lys Ala Phe Tyr Gln Val Leu His Leu Lys Gly Ile
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 Thr Glu Glu Gln Arg Asn Gln Tyr Ile Lys Thr Leu Arg Glu His Pro
 65 70 75 80
 Glu Arg Ala Gln Glu Val Phe Ser Glu Ser Leu Lys Asp Ser Lys Asn
 85 90 95
 Pro Asp Arg Arg Val Ala Gln Gln Asn Ala Phe Tyr Asn Val Leu Lys
 100 105 110
 Asn Asp Asn Leu Thr Glu Gln Glu Lys Asn Asn Tyr Ile Ala Gln Ile
 115 120 125
 Lys Glu Asn Pro Asp Arg Ser Gln Gln Val Trp Val Glu Ser Val Gln
 130 135 140
 Ser Ser Lys Ala Lys Glu Arg Gln Asn Ile Glu Asn Ala Asp Lys Ala
 145 150 155 160
 Ile Lys Asp Phe Gln Asp Asn Lys Ala Pro His Asp Lys Ser Ala Ala
 165 170 175
 Tyr Glu Ala Asn Ser Lys Leu Pro Lys Asp Leu Arg Asp Lys Asn Asn
 180 185 190
 Arg Phe Val Glu Lys Val Ser Ile Glu Lys Ala Ile Val Arg His Asp
 195 200 205
 Glu Arg Val Lys Ser Ala Asn Asp Ala Ile Ser Lys Leu Asn Glu Lys
 210 215 220
 Asp Ser Ile Glu Asn Arg Arg Leu Ala Gln Arg Glu Val Asn Lys Ala
 225 230 235 240
 Pro Met Asp Val Lys Glu His Leu Gln Lys Gln Leu Asp Ala Leu Val
 245 250 255
 Ala Gln Lys Asp Ala Glu Lys Lys Val Ala Pro Lys Val Glu Ala Pro

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5	Val Glu Val Pro Gln Ile	Gln Ser Pro Lys Val Glu Val Pro Gln Ser	
	Lys Leu Leu Gly Tyr Tyr	Gln Ser Leu Lys Asp Ser Phe Asn Tyr Gly	
10	Tyr Lys Tyr Leu Thr Asp Thr Tyr Lys Ser Tyr Lys Glu Lys Tyr Asp		
	Thr Ala Lys Tyr Tyr Tyr Asn Thr Tyr Tyr Lys Tyr Lys Gly Ala Ile		
15	Asp Gln Thr Val Leu Thr Val Leu Gly Ser Gly Ser Lys Ser Tyr Ile		
	Gln Pro Leu Lys Val Asp Asp Lys Asn Gly Tyr Leu Ala Lys Ser Tyr		
20	Ala Gln Val Arg Asn Tyr Val Thr Glu Ser Ile Asn Thr Gly Lys Val		
	Leu Tyr Thr Phe Tyr Gln Asn Pro Thr Leu Val Lys Thr Ala Ile Lys		
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	Ser Phe Trp Lys		
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	Thr Gly Tyr Ala Ala Gly Thr Gly His Gln Ala His Ala Ala Glu Val		
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	Gln Leu Asn Ala Ala Pro Ile Lys Asp Gly Ala Tyr Asp Ile His Phe		
45	Val Lys Asp Gly Phe Gln Tyr Asn Phe Thr Ser Asn Gly Thr Thr Trp		
	Ser Trp Ser Tyr Glu Ala Ala Asn Gly Gln Thr Ala Gly Phe Ser Asn		
50	Val Ala Gly Ala Asp Tyr Thr Thr Ser Tyr Asn Gln Gly Ser Asn Val		
	Gln Ser Val Ser Tyr Asn Ala Gln Ser Ser Asn Ser Asn Val Glu Ala		
55	Val Ser Ala Pro Thr Tyr His Asn Tyr Ser Thr Ser Thr Thr Ser Ser		
	Ser Val Arg Leu Ser Asn Gly Asn Thr Ala Gly Ala Thr Gly Ser Ser		

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Ala Ala Gln Ile Met Ala Gln Arg Thr Gly Val Ser Ala Ser Thr Trp
165 170 175

Ala Ala Ile Ile Ala Arg Glu Ser Asn Gly Gln Val Asn Ala Tyr Asn
5 180 185 190

Pro Ser Gly Ala Ser Gly Leu Phe Gln Thr Met Pro Gly Trp Gly Pro
195 200 205

Thr Asn Thr Val Asp Gln Gln Ile Asn Ala Ala Val Lys Ala Tyr Lys
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Ala Gln Gly Leu Gly Ala Trp Gly Phe
225 230

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<212> PRT
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<400> 47
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Leu Val Ile Ser Ile Thr Ala Gly Cys Gly Ile Gly Lys Glu Ala Glu
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Val Lys Lys Ser Phe Glu Lys Thr Leu Ser Met Tyr Pro Ile Lys Asn
25 35 40 45

Leu Glu Asp Leu Tyr Asp Lys Glu Gly Tyr Arg Asp Asp Gln Phe Asp
50 55 60

Lys Asn Asp Lys Gly Thr Trp Ile Ile Asn Ser Glu Met Val Ile Gln
30 65 70 75 80

Pro Asn Asn Glu Asp Met Val Ala Lys Gly Met Val Leu Tyr Met Asn
85 90 95

Arg Asn Thr Lys Thr Thr Asn Gly Tyr Tyr Tyr Val Asp Val Thr Lys
35 100 105 110

Asp Glu Asp Glu Gly Lys Pro His Asp Asn Glu Lys Arg Tyr Pro Val
115 120 125

Lys Met Val Asp Asn Lys Ile Ile Pro Thr Lys Glu Ile Lys Asp Glu
40 130 135 140

Lys Ile Lys Lys Glu Ile Glu Asn Phe Lys Phe Phe Val Gln Tyr Gly
145 150 155 160

Asp Phe Lys Asn Leu Lys Asn Tyr Lys Asp Gly Asp Ile Ser Tyr Asn
45 165 170 175

Pro Glu Val Pro Ser Tyr Ser Ala Lys Tyr Gln Leu Thr Asn Asp Asp
180 185 190

Tyr Asn Val Lys Gln Leu Arg Lys Arg Tyr Asp Ile Pro Thr Ser Lys
195 200 205

Ala Pro Lys Leu Leu Leu Lys Gly Ser Gly Asn Leu Lys Gly Ser Ser
210 215 220

Val Gly Tyr Lys Asp Ile Glu Phe Thr Phe Val Glu Lys Lys Glu Glu
225 230 235 240

Asn Ile Tyr Phe Ser Asp Ser Leu Asp Tyr Lys Lys Ser Gly Asp Val
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 <211> 514
 <212> PRT
 <213> Staphylococcus aureus

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Val Ser Leu Ser Ala Leu Pro Gln Ser Leu Ala Ile Thr His Glu Ser
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10 Gln Pro Thr Lys Gln Gln Arg Thr Val Leu Phe Asp Arg Ser His Gly
 35 40 45

Gln Thr Ala Gly Ala Ala Asp Trp Val Ser Asp Gly Ala Phe Ser Asp
 50 55 60

15 Tyr Ala Asp Ser Ile Gln Lys Gln Gly Tyr Asp Val Lys Ala Ile Asp
 65 70 75 80

Gly His Ser Asn Ile Thr Glu Ala Ser Leu Lys Ser Ser Lys Ile Phe
 85 90 95

20 Val Ile Pro Glu Ala Asn Ile Pro Phe Lys Glu Ser Glu Gln Ala Ala
 100 105 110

Ile Val Lys Tyr Val Lys Gln Gly Gly Asn Val Val Phe Ile Ser Asp
 115 120 125

25 His Tyr Asn Ala Asp Arg Asn Leu Asn Arg Ile Asp Ser Ser Glu Ala
 130 135 140

Met Asn Gly Tyr Arg Arg Gly Ala Tyr Glu Asp Met Ser Lys Gly Met
 145 150 155 160

30 Asn Ala Glu Glu Lys Ser Ser Thr Ala Met Gln Gly Val Lys Ser Ser
 165 170 175

Asp Trp Leu Ser Thr Asn Phe Gly Val Arg Phe Arg Tyr Asn Ala Leu
 180 185 190

35 Gly Asp Leu Asn Thr Ser Asn Ile Val Ser Ser Lys Glu Ser Phe Gly
 195 200 205

Ile Thr Glu Gly Val Lys Ser Val Ser Met His Ala Gly Ser Thr Leu
 210 215 220

40 Ala Ile Thr Asn Pro Glu Lys Ala Lys Gly Ile Val Tyr Thr Pro Glu
 225 230 235 240

Gln Leu Pro Ala Lys Ser Lys Trp Ser His Ala Val Asp Gln Gly Ile
 245 250 255

45 Tyr Asn Gly Gly Gly Lys Ala Glu Gly Pro Tyr Val Ala Ile Ser Lys
 260 265 270

Val Gly Lys Gly Lys Ala Ala Phe Ile Gly Asp Ser Ser Leu Val Glu
 275 280 285

50 Asp Ser Ser Pro Lys Tyr Val Arg Glu Asp Asn Gly Glu Lys Lys Lys
 290 295 300

Thr Tyr Asp Gly Phe Lys Glu Gln Asp Asn Gly Lys Leu Leu Asn Asn
 305 310 315 320

55 Ile Thr Ala Trp Met Ser Lys Asp Asn Asp Gly Lys Ser Leu Lys Ala
 325 330 335

Ser Ser Leu Thr Leu Asp Thr Lys Thr Lys Leu Leu Asp Phe Glu Arg

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5																	
10																	
15																	
20																	
25																	
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40																	
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55																	

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5 Gln Ser Leu Thr Phe His Ser Ala Phe Glu Ile Leu Gln Gln Tyr Leu
 165 170 175
 Pro Glu Met Lys Arg His Ala Asp Ile Ile Val Val Cys Tyr His Gly
 180 185 190
 Gly Phe Glu Lys Asp Leu Glu Ser Gly Thr Pro Thr Glu Val Leu Thr
 195 200 205
 10 Gly Glu Asn Glu Gly Tyr Ala Met Leu Glu Ala Phe Ser Lys Asp Ile
 210 215 220
 Asp Ile Phe Ile Thr Gly His Gln His Arg Gln Ile Ala Glu Arg Phe
 225 230 235 240
 15 Lys Gln Thr Ala Val Ile Gln Pro Gly Thr Arg Gly Thr Thr Val Gly
 245 250 255
 Arg Val Val Leu Ser Thr Asp Glu Tyr Glu Asn Leu Ser Val Glu Ser
 260 265 270
 20 Cys Glu Leu Leu Pro Val Ile Asp Asp Ser Thr Phe Thr Ile Asp Glu
 275 280 285
 Asp Asp Gln His Leu Arg Lys Gln Leu Glu Asp Trp Leu Asp Tyr Glu
 290 295 300
 25 Ile Thr Thr Leu Pro Tyr Asp Met Thr Ile Asn His Ala Phe Glu Ala
 305 310 315 320
 Arg Val Ala Pro His Pro Phe Thr Asn Phe Met Asn Tyr Ala Leu Leu
 325 330 335
 30 Glu Lys Ser Asp Ala Asp Val Ala Cys Thr Ala Leu Phe Asp Ser Ala
 340 345 350
 Ser Gly Phe Lys Gln Val Val Thr Met Arg Asp Val Ile Asn Asn Tyr
 355 360 365
 35 Pro Phe Pro Asn Thr Phe Lys Val Leu Ala Val Ser Gly Ala Lys Leu
 370 375 380
 Lys Glu Ala Ile Glu Arg Ser Ala Glu Tyr Phe Asp Val Lys Asn Asp
 385 390 395 400
 40 Glu Val Ser Val Ser Ala Asp Phe Leu Glu Pro Lys Pro Gln His Phe
 405 410 415
 Asn Tyr Asp Ile Tyr Gly Gly Val Ser Tyr Thr Ile His Val Gly Arg
 420 425 430
 45 Pro Lys Gly Gln Arg Val Ser Asn Met Met Ile Gln Gly His Ala Val
 435 440 445
 Asp Leu Lys Gln Thr Tyr Thr Ile Cys Val Asn Asn Tyr Arg Ala Val
 450 455 460
 50 Gly Gly Gly Gln Tyr Asp Met Tyr Ile Asp Ala Pro Val Val Lys Asp
 465 470 475 480
 Ile Gln Val Glu Gly Ala Gln Leu Leu Ile Asp Phe Leu Ser Asn Asn
 485 490 495
 55 Asn Leu Met Arg Ile Pro Gln Val Val Asp Phe Lys Val Glu Lys
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 <211> 324
 <212> PRT

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<213> Staphylococcus aureus

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Asn Gln Gln Val Ile Lys Ile Gly Tyr Leu Pro Ile Thr His Ser Ala
 35 40 45

Asn Leu Met Met Thr Lys Lys Leu Leu Ser Gln Tyr Asn His Pro Lys
 50 55 60

Tyr Lys Leu Glu Leu Val Lys Phe Asn Asn Trp Pro Asp Leu Met Asp
 65 70 75 80

Ala Leu Asn Ser Gly Arg Ile Asp Gly Ala Ser Thr Leu Ile Glu Leu
 85 90 95

Ala Met Lys Ser Lys Gln Lys Gly Ser Asn Ile Lys Ala Val Ala Leu
 100 105 110

Gly His His Glu Gly Asn Val Ile Met Gly Gln Lys Gly Met His Leu
 115 120 125

Asn Glu Phe Asn Asn Asn Gly Asp Asp Tyr His Phe Gly Ile Pro His
 130 135 140

Arg Tyr Ser Thr His Tyr Leu Leu Leu Glu Glu Leu Arg Lys Gln Leu
 145 150 155 160

Lys Ile Lys Pro Gly His Phe Ser Tyr His Glu Met Ser Pro Ala Glu
 165 170 175

Met Pro Ala Ala Leu Ser Glu His Arg Ile Thr Gly Tyr Ser Val Ala
 180 185 190

Glu Pro Phe Gly Ala Leu Gly Glu Lys Leu Gly Lys Gly Lys Thr Leu
 195 200 205

Lys His Gly Asp Asp Val Ile Pro Asp Ala Tyr Cys Cys Val Leu Val
 210 215 220

Leu Arg Gly Glu Leu Leu Asp Gln His Lys Asp Val Ala Gln Ala Phe
 225 230 235 240

Val Gln Asp Tyr Lys Lys Ser Gly Phe Lys Met Asn Asp Arg Lys Gln
 245 250 255

Ser Val Asp Ile Met Thr His His Phe Lys Gln Ser Arg Asp Val Leu
 260 265 270

Thr Gln Ser Ala Ala Trp Thr Ser Tyr Gly Asp Leu Thr Ile Lys Pro
 275 280 285

Ser Gly Tyr Gln Glu Ile Thr Thr Leu Val Lys Gln His His Leu Phe
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Asn Pro Pro Ala Tyr Asp Asp Phe Val Glu Pro Ser Leu Tyr Lys Glu
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 35 40 45
 10 Ala Ser Tyr Glu Asn Asp Met Asn Ile Val Lys Asp Gln Leu Glu Asn
 50 55 60
 Ala Gly Phe Asn Val Lys Met Asn Ile Gln Pro Asp Tyr Gly Ser Tyr
 65 70 75 80
 15 Arg Thr Gln Arg Gln Ala Gly Asn Tyr Asp Ile Gln Ile Asp Asp Trp
 85 90 95
 Met Thr Val Phe Gly Asp Pro Asn Tyr Ala Met Thr Ala Leu Phe Ser
 100 105 110
 20 Ser Thr Gly Ser Asn Ser Leu Leu Lys Asp Lys His Val Asp Gln Leu
 115 120 125
 Leu Asn Lys Ala Ser Thr Gln Asn Glu Ala Asp Val Lys Gln Thr Tyr
 130 135 140
 25 Lys Gln Ile Glu Asp Glu Val Val Phe Asp Lys Gly Tyr Met Ala Pro
 145 150 155 160
 Leu Tyr Gly Ser Lys Lys Asn Leu Val Tyr Asp Asn Lys Val Leu Asp
 165 170 175
 30 Lys Asn Ser Val Gly Leu Pro Asn Ser Arg Ala Leu Ile Trp Gln Gln
 180 185 190
 Phe Asp Tyr Asn Asn Ser Arg Glu Arg Asp Thr Arg Pro Leu Val Met
 195 200 205
 35 Thr Gln Gln Asp Gly Glu Ile Pro Thr Leu Asp Pro Ile Arg Ser Ile
 210 215 220
 Ala Pro Ser Val Tyr Ser Ile Asn Met Asn Met Tyr Thr Arg Leu Leu
 225 230 235 240
 40 Leu Leu Asp Glu Asn Asp His Leu Thr Thr Lys Gly Ser Leu Ser His
 245 250 255
 Asp Tyr Ala Val Asn Lys Asp Asn Lys Ala Phe Tyr Phe Leu Leu Arg
 260 265 270
 45 Asp Asp Asp Tyr Phe Ala Lys Val Val Asn Gly Gln Ala Arg Asn Thr
 275 280 285
 Gly Glu Arg Val Ser Ala Glu Asp Val Lys Phe Ser Leu Asp Arg Ala
 290 295 300
 50 Arg Asp Lys Lys Ser Val Pro Asn Asn Asn Thr Tyr Asn Met His Lys
 305 310 315 320
 His Ile Asn Asp Ile Lys Ile Leu Lys Asp Glu Asp Ile Asp Gln Leu
 325 330 335
 Arg Lys Glu Lys Asp Lys Asp Asp Lys Ser Ile Tyr Asp Lys Leu Ile
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 55 Lys Ala Tyr Asn Val Lys Ser Leu Thr Thr Asp Gly Gln Lys Val Asn
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 5 Met Pro Arg Glu Val Asn Tyr Leu Thr His Ser Ser Ala Gly Ile Leu
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 10 Gly Asp Ser Ser Thr Ile Pro Ala Asn Ser Asp Gly Lys Asn Ala Leu
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 Asp Val Asn Gln Lys His Phe Asp Leu Ile Lys Ser Asp Lys Asn Leu
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 5 Arg Tyr Ala Lys Leu Ser Leu Glu Asp Leu Ile Asn Pro Ala Ile Glu
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 10 Arg Gln Gln His Ala Arg Leu Thr Lys Tyr His Glu Thr Ala Gln Val
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 5 Ser Met His His Arg Val Ser Asp Val His Gln Phe Gln Asn His Ala
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 195 200 205
 10 Lys Tyr Tyr Ala Gly Leu Ser Tyr Glu Gln Asp Ala Arg Thr Trp Lys
 210 215 220
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 225 230 235 240
 15 Leu Asn Val Asp Gly Ile Ala Leu Val Lys Asn Ala His Pro His Pro
 245 250 255
 Lys Arg Lys Lys Leu Val Gln Tyr Leu Thr Ser Arg Ser Val Gln Gln
 260 265 270
 20 Arg Leu Val Ala Glu Phe Asp Ala Lys Ser Ile Arg Lys Asp Val Ser
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 5 Pro Lys Leu Asp His Ile Thr Val Thr Tyr Gln Glu Asp Gly Asn Asn
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 10 Pro Val Asn Lys Val Gln Asp Ile Glu Asn Asn Gln Asn Leu Lys Val
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 Ser Lys Glu Ser Gly Phe Arg Thr Ser Leu Leu Met Tyr Asn His Thr
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 15 Asn Lys Lys Met Thr Lys Ser Val Arg Glu Ala Leu Asp His Ile Ile
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 Asp Arg Gln Gly Ile Ala Asp His Ile Tyr Gln Gly Tyr Ala Lys Pro
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 30 Gly Tyr Leu Lys Asp Arg Ser Ala Trp Asp Ala Thr Met Tyr Ser Phe
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 370 375 380
 35 Lys Lys Asp Gly Ala Ile Asn Lys Gly Asp Tyr Asn Asn Ser Asn Val
 385 390 395 400
 Asp Asp Leu Ile Asn Gln Leu Asn His Thr Val Asp Val Lys Glu Arg
 405 410 415
 40 His Asn Ile Ser Asn Asp Ile Ile Lys Leu Ser Ser Arg Asp Val Pro
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 Asn Ser Tyr Ile Ala Tyr Asn Asp Gln Ile Val Ala Ala Asn Ser Lys
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 5 Lys Met Lys Asp Asn Ser Phe Leu Ser Ser Leu Glu Leu Ser Ala Asp
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 Ala Ser Asp Glu Ile Val Lys Gly Leu Gly Ile Pro Lys Ser Val Val
 85 90 95
 10 Asn Ala Ser Lys Ile Lys Met Ser Tyr Gly His Asp Pro Lys Lys Glu
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 115 120 125
 15 Lys Phe Gln Leu Ala Ala Asp Lys Asp Lys His Tyr Phe Glu Ser Pro
 130 135 140
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 145 150 155 160
 20 Tyr Ser Lys Leu Thr Gly Glu Asp Glu Glu Thr Ala Lys Glu Asn Gly
 165 170 175
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 180 185 190
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 210 215 220
 30 Glu Ile Lys Val Asn Gly Glu Lys Tyr Lys Val Arg Pro Val Thr Leu
 225 230 235 240
 Thr Leu Ser Arg Ala Asp Thr Lys Lys Ile Thr Leu Ala Val Leu Glu
 245 250 255
 35 Glu Ala Lys Lys Asp Lys Asp Leu Lys Lys Leu Met Glu Glu Gln Gly
 260 265 270
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 40 Lys Glu Thr Lys Lys Asp Glu Phe Ala Lys Ile Gln Ser Lys Ile Tyr
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 Thr Ile Lys Thr Asp Ile Lys Gln Glu Asp Ser Asn Thr His Val Tyr
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 25 Lys Asp Gly Lys Val Ile Val Ile Gly Ile Gln Leu Tyr Lys Asp Arg
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 Glu Lys Met Tyr Tyr Phe Ala Tyr Glu Ile Lys Asp Gly Lys Ala Glu
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Trp His Glu Trp Val Gln Ala Ala Lys Ala Lys Pro Val Tyr Gly Ala
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 5 Lys Glu Phe Leu Lys Tyr Ala Asp Lys Lys Gly Val Asp Ile Tyr Tyr
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 10 Leu Lys Gln Gln Gly Ile Pro Gln Ala Lys Lys Ser His Ile Leu Leu
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 15 Lys Asp His Lys Leu Val Met Leu Phe Gly Asp Asn Leu Leu Asp Phe
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 Thr Asp Pro Lys Glu Ala Thr Ala Glu Ser Arg Glu Ala Leu Ile Glu
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 85 90 95
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 Ile Pro Asn His Asp Lys Ala Ala Pro Thr Ser Thr Thr Pro Pro Ser
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 Gly His Glu Arg Tyr Gly Lys Thr Tyr Lys Gly Ile Met Pro Asn Trp
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 Glu Pro Gly Lys Lys Val His Leu Val Gly His Ser Met Gly Gly Gln
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 Thr Ile Arg Leu Met Glu Glu Phe Leu Arg Asn Gly Asn Lys Glu Glu
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 435 440 445
 Gly Gly His Asn Asn Met Val Ala Ser Ile Thr Thr Leu Ala Thr Pro
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20	Lys	Arg	Lys	Gly 660	Ala	Glu	Leu	Ala	Asn 665	Phe	Tyr	Thr	Gly	Ile 670	Ile	Asn	
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	Ser	Thr	Gly 35	Ile	His	Ala	Pro	Lys 40	Ser	Ala	Lys	Lys	Leu 45	Glu	Thr	Lys	
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	Asn 130	Asn	Ile	Pro	Lys	Glu	Tyr 135	Lys	Lys	Pro	Val	Glu 140	Leu	Met	Met	Asn	
55	Tyr 145	Phe	Lys	Ala	Leu	Asn 150	Ser	Thr	Ile	Ala	Asn 155	Val	Asp	Glu	Asp	Ile 160	
	Glu	Lys	Leu	Ser	Tyr 165	Gln	Pro	Gln	Asn	Lys 170	Ile	Asn	Val	Val	Asp 175	Val	

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 Ile Thr Asn Glu Asn His Ala Asn Lys Asp Lys Ile Asn Ile Lys Asn
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 Lys Asn Val Arg Ile Tyr Gln Asn Leu Thr Tyr Asn Arg Val Phe Pro
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 Asn Ser Lys Leu Asp Ile Ile Thr Pro Val Asp Met Ser Ser Asn Ala
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 Lys Leu Pro Val Ile Phe Trp Met His Gly Gly Gly Tyr Ile Ala Gly
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 Asp Lys Gln Tyr Lys Asn Pro Leu Leu Ala Lys Ile Ala Glu Gln Gly
 100 105 110
 Tyr Ile Val Val Asn Val Asn Tyr Ala Leu Ala Pro Gln Tyr Lys Tyr
 115 120 125
 Pro Thr Pro Leu Ile Gln Met Asn Gln Ala Thr Gln Phe Ile Lys Glu
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 Asp Ser Ala Gly Ala Gln Leu Ala Ser Gln Phe Thr Ala Ile Gln Thr
 165 170 175
 Asn Asp Arg Leu Arg Glu Ala Met Lys Phe Asp Gln Ser Phe Lys Pro
 180 185 190
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 195 200 205
 Thr Val Arg Glu Thr Glu Phe Pro Arg Ile Gln Leu Phe Met Lys Ser
 210 215 220
 Tyr Thr Gly Glu Glu Asp Trp Glu Lys Ser Phe Lys Asn Ile Ser Gln
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 Met Ser Thr Val Lys Gln Ser Thr Lys Asn Tyr Pro Pro Thr Phe Leu
 245 250 255
 Ser Val Gly Asp Ser Asp Pro Phe Glu Ser Gln Asn Ile Glu Phe Ser
 260 265 270
 Lys Lys Leu Gln Glu Leu Asn Val Pro Val Asp Thr Leu Phe Tyr Asp
 275 280 285
 Gly Thr His His Leu His His Gln Tyr Gln Phe His Leu Asn Lys Pro
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 Glu Ser Ile Asp Asn Ile Lys Lys Val Leu Leu Phe Leu Ser Arg Asn
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 10 Glu Ser Ile Lys Gln Asn Asn Gln Leu Asp Asn Asn Leu Val Phe Pro
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 40 325 330 335
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 Ile Leu Met Gly Ile Ile Ile Leu Ile Val Ile Leu Asn Ala Ile Phe
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5 Ser Val Val Thr Val Glu Asn Glu Thr Ser Lys Asp Ser Ser Leu Pro
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 20 25 30
 Leu Thr Met Thr Leu Asp Lys Leu Asp Leu Pro Ile Val Gly Lys Pro
 35 40 45
 Thr Ser Tyr Lys Thr Leu Pro Asn Arg Tyr Lys Asp Val Pro Glu Ile
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 30 Gly Gln Pro Met Glu Pro Asn Val Glu Ala Val Lys Lys Leu Lys Pro
 65 70 75 80
 35 Thr His Val Leu Ser Val Ser Thr Ile Lys Asp Glu Met Gln Pro Phe
 85 90 95
 Tyr Lys Gln Leu Asn Met Lys Gly Tyr Phe Tyr Asp Phe Asp Ser Leu
 100 105 110
 40 Lys Gly Met Gln Lys Ser Ile Thr Gln Leu Gly Asp Gln Phe Asn Arg
 115 120 125
 Lys Ala Gln Ala Lys Glu Leu Asn Asp His Leu Asn Ser Val Lys Gln
 130 135 140
 45 Lys Ile Glu Asn Lys Ala Ala Lys Gln Lys Lys His Pro Lys Val Leu
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 50 Ile Leu Met Gly Val Pro Gly Ser Tyr Leu Val Ala Thr Asp Lys Ser
 165 170 175
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 180 185 190
 55 Val Lys Asp Arg Gln Tyr Ile Ser Ser Asn Thr Glu Asn Leu Leu Asn
 195 200 205
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 210 215 220
 Val Lys Lys Met Phe Gln Lys Glu Phe Lys Gln Asn Asp Ile Trp Lys
 225 230 235 240

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His Phe Lys Ala Val Lys Asn Asn His Val Tyr Asp Leu Glu Glu Val
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 Lys Gln Glu Arg Ala Asn Tyr Glu Lys Leu Gln Gln Lys Phe Gln Met
 35 40 45
 20 Leu Met Ser Lys His Gln Glu His Val Arg Pro Gln Phe Glu Ser Leu
 50 55 60
 Glu Lys Ile Asn Lys Asp Ile Val Gly Trp Ile Lys Leu Ser Gly Thr
 65 70 75 80
 25 Ser Leu Asn Tyr Pro Val Leu Gln Gly Lys Thr Asn His Asp Tyr Leu
 85 90 95
 Asn Leu Asp Phe Glu Arg Glu His Arg Arg Lys Gly Ser Ile Phe Met
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 30 Asp Phe Arg Asn Glu Leu Lys Asn Leu Asn His Asn Thr Ile Leu Tyr
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 Lys Tyr Gly Lys Tyr Gln Leu Gln Val Phe Ser Ala Tyr Lys Thr Thr
 165 170 175
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 195 200 205
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	Ser	Ser	Glu	Ala	Lys	Ala	Tyr	Tyr	Ile	Asn	Gln	Asn	Glu	Thr	Asn	Val
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Lys Asp Gln Glu Thr Ile Asp Ser Phe Ala Leu Ser Gln Glu Thr Phe
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 5 Thr Gly Gly Val Thr Pro Ala Ala Thr Ser Asn Asp Lys His Tyr Lys
 130 135 140
 Leu Asn Val Thr Tyr Lys Asp Lys Ala Glu Thr Phe Thr Gly Gly Phe
 145 150 155 160
 10 Pro Val Tyr Glu Gly Asn Lys Pro Val Leu Thr Leu Lys Glu Leu Asp
 165 170 175
 Phe Arg Ile Arg Gln Thr Leu Ile Lys Ser Lys Lys Leu Tyr Asn Asn
 180 185 190
 15 Ser Tyr Asn Lys Gly Gln Ile Lys Ile Thr Gly Ala Asp Asn Asn Tyr
 195 200 205
 Thr Ile Asp Leu Ser Lys Arg Leu Pro Ser Thr Asp Ala Asn Arg Tyr
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 Leu Ser Ile His Pro Asn Phe Ser Arg Leu Gln Leu Thr Thr Lys Lys
 50 55 60
 40 Tyr Asp Asn Pro Phe Asn Pro Pro Met Phe Ala Arg Val Phe Arg Lys
 65 70 75 80
 His Leu Glu Gly Gly Ile Ile Glu Ser Ile Lys Gln Ile Gly Asn Asp
 85 90 95
 45 Arg Arg Ile Glu Ile Asp Ile Lys Ser Lys Asp Glu Ile Gly Asp Thr
 100 105 110
 Ile Tyr Arg Thr Val Ile Leu Glu Ile Met Gly Lys His Ser Asn Leu
 115 120 125
 50 Ile Leu Val Asp Glu Asn Arg Lys Ile Ile Glu Gly Phe Lys His Leu
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 Thr Pro Asn Thr Asn His Tyr Arg Thr Val Met Pro Gly Phe Asn Tyr
 145 150 155 160
 55 Glu Ala Pro Pro Thr Gln His Lys Ile Asn Pro Tyr Asp Ile Thr Gly
 165 170 175
 Ala Glu Val Leu Lys Tyr Ile Asp Phe Asn Ala Gly Asn Ile Ala Lys
 180 185 190

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Gln Leu Leu Asn Gln Phe Glu Gly Phe Ser Pro Leu Ile Thr Asn Glu
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 5 Ile Val Ser Arg Arg Gln Phe Met Thr Ser Ser Thr Leu Pro Glu Ala
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 Phe Asp Glu Val Met Ala Glu Thr Lys Leu Pro Pro Thr Pro Ile Phe
 225 230 235
 10 His Lys Asn His Glu Thr Gly Lys Glu Asp Phe Tyr Phe Ile Lys Leu
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 Asn Gln Phe Asn Asp Asp Thr Val Thr Tyr Asp Ser Leu Asn Asp Leu
 260 265
 15 Leu Asp Arg Phe Tyr Asp Ala Arg Gly Glu Arg Glu Arg Val Lys Gln
 275 280 285
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 290 295 300
 20 Gln Asn Lys Leu Ala Lys Leu Ile Glu Glu Tyr Glu Gln Ser Lys Asn
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 Lys Asp Thr Glu Gln Leu Tyr Gly Glu Leu Ile Thr Ala Asn Ile Tyr
 325 330 335
 Arg Ile Lys Gln Gly Asp Lys Glu Val Thr Ala Leu Asn Tyr Tyr Thr
 340 345 350
 25 Asn Glu Glu Val Val Ile Pro Leu Asn Pro Thr Lys Ser Pro Ser Ala
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 Asn Ala Gln Tyr Tyr Tyr Lys Gln Tyr Asn Arg Met Lys Thr Arg Glu
 370 375 380
 30 Arg Glu Leu Gln His Gln Ile Gln Leu Thr Lys Asp Asn Ile Asp Tyr
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 405 410 415
 35 Asp Glu Ile Arg Asp Glu Leu Ala Glu Gln Gly Phe Met Lys Gln Arg
 420 425 430
 Lys Asn Gln Thr Lys Lys Lys Lys Ala Gln Ile Gln Leu Gln His Tyr
 435 440 445
 40 Val Ser Thr Asp Gly Asp Asp Ile Tyr Val Gly Lys Asn Asn Lys Gln
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 45 His Thr Lys Asp Ile Pro Gly Ser His Val Val Ile Phe Asn Asp Ala
 485 490 495
 Pro Ser Asp Thr Thr Ile Lys Glu Ala Ala Met Leu Ala Gly Tyr Phe
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 50 Ser Lys Ala Gly Asn Ser Gly Gln Ile Pro Val Asp Tyr Thr Leu Ile
 515 520 525
 Lys Asn Val His Lys Pro Ser Gly Ala Lys Pro Gly Phe Val Thr Tyr
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 Lys Met Lys Gln Ser

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565

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 <212> PRT
 <213> Staphylococcus aureus

 <400> 78
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 Asp Pro Asp Leu Glu Lys Tyr Glu Glu Ile Glu Lys Lys Met Lys Gly
 50 55 60
 Ile Lys Asp Ala Pro Ser Leu Asp Lys Leu Asp Pro Leu Met Thr Glu
 65 70 75 80
 Lys Ser Phe Thr Asn Ser Lys Gly Ile Gln Gly Trp Lys Asp Tyr Lys
 85 90 95
 Glu Leu Met Gly Lys Val Glu Leu Ala Asp Tyr Arg Phe Thr Lys Asp
 100 105 110
 Ser Lys Gly Ser Ser Ile Lys Asp Val Asp Ala Phe Phe Lys Gly Lys
 115 120 125
 Lys Gly Ile Lys Arg Lys Val Ile Glu Thr His Asp Asp Val Lys Gln
 130 135 140
 Val Asp Tyr Trp Tyr Val Asp Pro Asp Gly Lys Lys Ile Gly Asn Ser
 145 150 155 160
 Asn Thr Pro Val Phe Tyr Ala Glu Ile Met Thr Lys Tyr Lys Asp Gly
 165 170 175
 Lys Leu Val Tyr Ala Ser Val Glu Pro Gly Ser Tyr Val Ile His Lys
 180 185 190
 Asp Asp Ala Ile Lys Tyr Asp Asp Tyr Ser Lys Leu Lys Lys Leu Ser
 195 200 205
 Gln Leu Thr Lys Leu Asp His Pro Lys Pro Val Pro Tyr Ser Val Ala
 210 215 220
 Gln Ile Lys Ser Phe Gly Val Pro Leu Thr Ser Val Ser Phe Met Thr
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 His Gly Ser Lys Asp Thr Lys Asp Glu Val Leu Pro Ala Leu Ala Tyr
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 Phe Thr Phe Ser Pro Lys Asn Tyr Glu Asp Lys Ser Asn Pro Asp Pro
 260 265 270
 Lys Val Leu Asn Leu Val His Met Asp Phe Leu Asn Ala Ser Ser Asp
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<212> PRT
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<400> 79

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	Arg	Gly	Tyr	Glu	Gln	Asn	Val	Phe	Ala	Lys	Ser	Glu	Leu	Asn	Ala	Asn
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	Ser	Asp	Thr	Ser	Leu	Lys	Leu	Asp	Asn	Lys	Ser	Ala	Ile	Ser	Asn	Gly
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	Ile	Ala	Leu	Lys	Tyr	Lys	Thr	Thr	Val	Ser	Asn	Ile	Gln	Asn	Thr	Asn
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	Glu	Asn	Arg	Gly	Trp	Asp	Phe	Asp	Gly	Ser	Tyr	Gly	Trp	Gln	Cys	Phe
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45	Asp	Leu	Val	Asn	Val	Tyr	Trp	Asn	His	Leu	Tyr	Gly	His	Gly	Leu	Lys
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 Phe Lys Lys Ala
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 Pro Glu Glu Met Asp Ala Ser Lys Tyr Val Gly Gln Gly Phe Gln Pro
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 Pro Ala Glu Lys Asp Ala Ile Glu Phe Ala Lys Lys His Lys Asp Lys
 50 55 60
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 Ile Ala Lys Arg Gly Glu Gln Phe Phe Met Asp Asn Phe Gly Leu Lys
 65 70 75
 Val Lys Ala Thr Asn Val Ile Gly Ser Gly Asp Gly Val Glu Val Phe
 85 90 95
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 Val His Cys Asp Asp His Asp Ile Val Phe Asn Ala Ser Ile Pro Phe
 100 105
 Asp Lys Ser Ile Ile Asp Ser Asp Ser Ser Leu Arg Ser Lys Asp Lys
 115 120 125
 30
 Gly Asp Asp Met Ser Thr Leu Val Gly Ala Val Leu Ser Gly Phe Glu
 130 135 140
 Tyr Arg Ala Gln Lys Glu Lys Tyr Asp Lys Leu Tyr Lys Phe Phe Lys
 145 150 155
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 Asp Asn Glu Glu Lys Tyr Gln Tyr Thr Gly Phe Thr Lys Glu Ala Ile
 165 170 175
 Asn Lys Thr Gln Asn Ser Gly Tyr Glu Asn Glu Tyr Phe Tyr Ile Ser
 180 185 190
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 Ala Ile Pro Tyr Asn Leu Ala Glu Tyr Arg Asp Tyr Phe Glu Pro Leu
 195 200 205
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 210 215 220
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 Lys Gln Leu Lys Asp Lys Ser Lys Val Ser Val Thr Thr Thr Leu Phe
 225 230 235
 Ser Lys Lys Lys Asn Tyr Thr Lys Lys Ser Asn Ser Glu Asn Val Ile
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 Lys Met Ala Glu Glu Ile Lys Lys Asp Lys Glu Ile Pro Asn Gly Ile
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 35 40 45
 Gln Ile Ser Asp Ser Lys Ser Lys Ile Lys Gly Leu Glu Lys Asp Lys
 50 55 60
 15 Glu Asn Ser Lys Lys Thr Ala Ser Asn Asn Thr Lys Ile Lys Leu Met
 65 70 75 80
 Asn Val Thr Ser Thr Tyr Tyr Asp Lys Val Ala Lys Ala Leu Lys Ser
 85 90 95
 20 Tyr Asn Asp Ile Glu Lys Asp Val Ser Lys Asn Lys Gly Asp Lys Asn
 100 105 110
 Val Gln Ser Lys Leu Asn Gln Ile Ser Asn Asp Ile Gln Ser Ala His
 115 120 125
 25 Thr Ser Tyr Lys Asp Ala Ile Asp Gly Leu Ser Leu Ser Asp Asp Asp
 130 135 140
 Lys Lys Thr Ser Lys Asn Ile Asp Lys Leu Asn Ser Asp Leu Asn His
 145 150 155 160
 30 Ala Phe Asp Asp Ile Lys Asn Gly Tyr Gln Asn Lys Asp Lys Lys Gln
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 50 Ile Leu Leu Ser Ala Cys Ser Met Met Phe Ser Asn Asn Asp Asn Ser
 50 55 60
 Thr Asn Asn Gln Ser Ser Lys Thr Gln Leu Thr Gln Lys Asp Glu Asn
 65 70 75 80
 Lys Asn Glu Asp Lys Pro Glu Glu Lys Ser Glu Thr Ala Thr Asp Glu
 85 90 95
 55 Asp Leu Gln Ser Thr Glu Glu Val Pro Ala Asn Glu Asn Thr Glu Asn
 100 105 110

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Asn Gln His Glu Ile Asp Glu Ile Thr Thr Lys Asp Gln Ser Asp Asp
 115 120 125
 5 Asp Ile Asn Thr Pro Asn Val Ala Glu Asp Lys Ser Gln Asp Asp Leu
 130 135 140
 Lys Asp Asp Leu Lys Glu Lys Gln Gln Ser Ser Asn His His Gln Ser
 145 150 155 160
 10 Thr Gln Pro Lys Thr Ser Pro Ser Thr Glu Thr Asn Thr Gln Gln Ser
 165 170 175
 Phe Ala Asn Cys Lys Gln Leu Arg Gln Val Tyr Pro Asn Gly Val Thr
 180 185 190
 15 Ala Asp His Pro Ala Tyr Arg Pro His Leu Asp Arg Asp Lys Asp Lys
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 Arg Ala Cys Glu Pro Asp Lys Tyr
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 <213> Staphylococcus aureus
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 30 Thr Glu Lys Gln Met Thr Pro Gln Glu Ala Glu Asp Ile Val Arg Asn
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 Asp Tyr Lys Ala Arg Gly Val Asn Glu Tyr Gln Thr Leu Asn Tyr Lys
 65 70 75 80
 35 Thr Asn Leu Glu Arg Ser Asn Gln His Glu Tyr Tyr Val Glu His Leu
 85 90 95
 Val Arg Asp Ala Val Gly Thr Pro Leu Lys Arg Cys Ala Ile Val Asn
 100 105 110
 40 Arg His Asn Gly Thr Ile Ile Asn Ile Phe Asp Asp Met Ser Glu Lys
 115 120 125
 Asp Lys Glu Glu Phe Glu Ala Phe Lys Lys Arg Ser Pro Lys Tyr Asn
 130 135 140
 45 Pro Gly Met Asn Asn His Asp Glu Thr Asp Gly Glu Ser Glu Asp Ile
 145 150 155 160
 Gln His His Asp Ile Asp Asn Asn Lys Ala Ile Gln Asn Asp Ile Pro
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 50 Asp Gln Lys Val Asp Asp Lys Asn Asp Lys Asn Ala Val Asn Lys Glu
 180 185 190
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 55 <210> 84
 <211> 457
 <212> PRT

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<213> Staphylococcus aureus

<400> 84

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10 Tyr Ser Val Glu Tyr Lys Asn Thr Ala Thr Phe Asn Lys Leu Val Lys
35 40 45

Lys Lys Ser Leu Asn Val Val Tyr Asn Ile Pro Glu Leu His Val Ala
50 55 60

15 Gln Ile Lys Met Thr Lys Met His Ala Asn Ala Leu Ala Asn Tyr Lys
65 70 75 80

Asn Asp Ile Lys Tyr Ile Asn Ala Thr Cys Ser Thr Cys Ile Thr Ser
85 90 95

20 Glu Lys Thr Ile Asp Arg Thr Ser Asn Glu Ser Leu Phe Ser Arg Gln
100 105 110

Trp Asp Met Asn Lys Ile Thr Asn Asn Gly Ala Ser Tyr Asp Asp Leu
115 120 125

25 Pro Lys His Ala Asn Thr Lys Ile Ala Ile Ile Asp Thr Gly Val Met
130 135 140

Lys Asn His Asp Asp Leu Lys Asn Asn Phe Ser Thr Asp Ser Lys Asn
145 150 155 160

Leu Val Pro Leu Asn Gly Phe Arg Gly Thr Glu Pro Glu Glu Thr Gly
165 170 175

30 Asp Val His Asp Val Asn Asp Arg Lys Gly His Gly Thr Met Val Ser
180 185 190

Gly Gln Thr Ser Ala Asn Gly Lys Leu Ile Gly Val Ala Pro Asn Asn
195 200 205

35 Lys Phe Thr Met Tyr Arg Val Phe Gly Ser Lys Lys Thr Glu Leu Leu
210 215 220

Trp Val Ser Lys Ala Ile Val Gln Ala Ala Asn Asp Gly Asn Gln Val
225 230 235 240

40 Ile Asn Ile Ser Val Gly Ser Tyr Ile Ile Leu Asp Lys Asn Asp His
245 250 255

Gln Thr Phe Arg Lys Asp Glu Lys Val Glu Tyr Asp Ala Leu Gln Lys
260 265 270

45 Ala Ile Asn Tyr Ala Lys Lys Lys Lys Ser Ile Val Val Ala Ala Ala
275 280 285

Gly Asn Asp Gly Ile Asp Val Asn Asp Lys Gln Lys Leu Lys Leu Gln
290 295 300

50 Arg Glu Tyr Gln Gly Asn Gly Glu Val Lys Asp Val Pro Ala Ser Met
305 310 315 320

Asp Asn Val Val Thr Val Gly Ser Thr Asp Gln Lys Ser Asn Leu Ser
325 330 335

55 Glu Phe Ser Asn Phe Gly Met Asn Tyr Thr Asp Ile Ala Ala Pro Gly
340 345 350

Gly Ser Phe Ala Tyr Leu Asn Gln Phe Gly Val Asp Lys Trp Met Asn

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	Glu	Gly	Tyr	Met	His	Lys	Glu	Asn	Ile	Leu	Thr	Thr	Ala	Asn	Asn	Gly	
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5	Arg	Tyr	Ile	Tyr	Gln	Ala	Gly	Thr	Ser	Leu	Ala	Thr	Pro	Lys	Val	Ser	
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	Lys	Pro	Phe	Ser	Arg	Tyr	Gly	His	Gly	Glu	Leu	Asp	Val	Tyr	Lys	Ala	
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	Leu	Ile	Ser	Ser	Lys	Ala	Gly	Asp	Val	Thr	Val	Ala	Asp	Thr	Met	Lys	
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30	Lys	Ile	Gly	Lys	Asp	Gln	Ile	Ala	Asn	Ala	Ser	Phe	Thr	Glu	Met	Leu	
		50					55					60					
	Asn	Lys	Ile	Leu	Ala	Asp	Lys	Tyr	Lys	Asn	Lys	Val	Asn	Asp	Lys	Lys	
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35	Ile	Asp	Glu	Gln	Ile	Glu	Lys	Met	Gln	Lys	Gln	Tyr	Gly	Gly	Lys	Asp	
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	Lys	Phe	Glu	Lys	Ala	Leu	Gln	Gln	Gln	Gly	Leu	Thr	Ala	Asp	Lys	Tyr	
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40	Lys	Glu	Asn	Leu	Arg	Thr	Ala	Ala	Tyr	His	Lys	Glu	Leu	Leu	Ser	Asp	
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	Lys	Ile	Lys	Ile	Ser	Asp	Ser	Glu	Ile	Lys	Glu	Asp	Ser	Lys	Lys	Ala	
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Ile Lys Ala Asp Lys₂₄₅ Pro Thr Asp Phe Asn₂₅₀ Ser Glu Lys Gln Ser Leu
 Lys Glu Lys Leu₂₆₀ Val Asp Gln Lys Val₂₆₅ Gln Lys Asn Pro Lys₂₇₀ Leu Leu
 Thr Asp Ala₂₇₅ Tyr Lys Asp Leu Leu₂₈₀ Lys Glu Tyr Asp Val₂₈₅ Asp Phe Lys
 Asp Arg₂₉₀ Asp Ile Lys Ser Val₂₉₅ Val Glu Asp Lys Ile₃₀₀ Leu Asn Pro Glu
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 Ala Lys Asp₃₅ Lys Lys His Val₄₀ Gln Val Asn Val Glu Asp₄₅ Lys Ser Val
 Pro Thr₅₀ Asp Val Arg Asn Leu₅₅ Ala Gln Lys Asp Tyr₆₀ Leu Ser Tyr Val
 Thr Ser Leu Asp Lys Ile₇₀ Tyr Asn Lys Glu Lys₇₅ Ala Ser Tyr Thr Leu₈₀
 Gly Glu Pro Phe Lys₈₅ Ile Tyr Lys Phe Asn₉₀ Lys Lys Ser Asp Gly₉₅ Asn
 Tyr Tyr Phe Pro₁₀₀ Val Leu Asn Thr Glu₁₀₅ Gly Asn Ile Asp Tyr₁₁₀ Ile Val
 Thr Ile Ser₁₁₅ Pro Lys Ile Thr Lys₁₂₀ Tyr Ser Ser Ser Ser₁₂₅ Ser Lys Tyr
 Thr Ile Asn Val Ser Pro Phe₁₃₅ Leu Ser Lys Val Leu₁₄₀ Asn Gln Tyr Lys
 Asp Gln Gln Ile Thr Ile₁₅₀ Leu Thr Asn Ser Lys₁₅₅ Gly Tyr Tyr Val Val₁₆₀
 Thr Gln Asn His Lys₁₆₅ Ala Lys Leu Val Leu₁₇₀ Lys Thr Pro Arg Leu Glu₁₇₅
 Asp Lys Lys Leu₁₈₀ Lys Lys Thr Glu Ser₁₈₅ Ile Pro Thr Gly Asn₁₉₀ Asn Val
 Thr Gln Leu Lys Gln Lys Ala Ser₂₀₀ Val Thr Met Pro Thr₂₀₅ Ser Gln Phe
 Lys Ser₂₁₀ Asn Asn Tyr Thr Tyr₂₁₅ Asn Glu Gln Tyr Ile₂₂₀ Asn Lys Leu Glu
 Asn Phe Lys Ile Arg Glu₂₃₀ Thr Gln Gly Asn Asn₂₃₅ Gly Trp Cys Ala Gly₂₄₀
 Tyr Thr Met Ser Glu₂₄₅ Leu Leu Asn Ala Thr₂₅₀ Tyr Asn Thr Asn Lys Tyr₂₅₅

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His Ala Glu Ala Val Met Arg Phe Leu His Pro Asn Leu Gln Gly Gln
 260 265 270
 Arg Phe Gln Phe Thr Gly Leu Thr Pro Arg Glu Met Ile Tyr Phe Gly
 275 280 285
 5 Gln Thr Gln Gly Arg Ser Pro Gln Leu Leu Asn Arg Met Thr Thr Tyr
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 10 Gly Ser Arg Val Glu Ser Arg Asn Gly Met His Ala Gly His Ala Met
 325 330 335
 Ala Val Val Gly Asn Ala Lys Leu Asp Asn Gly Gln Glu Val Ile Ile
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 40 Asp Lys Thr Leu Lys Ile Ser Asp Lys Arg Ser Lys Thr Arg Gly Tyr
 85 90 95
 Ala Ile Asp Met Asn Pro Phe His Glu Asn Lys Lys Thr Leu Thr Ile
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 115 120 125
 Gly Ser Val Arg Ile Ser Asp Val Asp Leu Glu Asn Thr Ser Ile Gln
 130 135 140
 Ser Ile Asn Gly Glu Val Val Ile Lys Asn Ser Asn Leu Asp Ala Leu
 145 150 155 160
 50 Asp Ser Lys Thr Asn Asn Ser Ser Thr Tyr Ile Ser Lys Ser Asn Ile
 165 170 175
 Lys Asn Ser Asn Ile Lys Val Val Ile Gly Thr Leu Gln Ile Asp Lys
 180 185 190
 55 Ser Gln Ile Lys Gln Ser Ile Phe Leu Asn Asp His Gly Asp Ile Glu

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10	Thr	Asn	Gly	Lys 260	Val	Gly	Lys	Ser	Asp 265	Asn	Val	Leu	Glu	Phe 270	Tyr	Thr
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	Lys	Arg 50	Val	Ala	Val	Leu	Thr 55	Gly	Phe	Tyr	Val	Gly 60	Asp	Phe	Ile	Lys
30	Leu 65	Gly	Ile	Lys	Pro	Ile 70	Ala	Val	Ser	Asp	Ile 75	Thr	Lys	Asp	Ser	Ser 80
	Ile	Leu	Lys	Pro	Tyr 85	Leu	Lys	Gly	Val	Asp 90	Tyr	Ile	Gly	Glu	Asn 95	Asp
35	Val	Glu	Arg	Val 100	Ala	Lys	Ala	Lys	Pro 105	Asp	Leu	Ile	Val	Val 110	Asp	Ala
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40	Tyr	Thr 130	Tyr	Asn	Lys	Tyr	Asn 135	His	Lys	Glu	Ile	Leu 140	Lys	Glu	Ile	Gly
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	Gln	Ala	Thr	Ala 180	Ser	Val	Phe	Glu	Pro 185	Asp	Glu	Lys	Gln	Ile 190	Tyr	Ile
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55	Lys 225	Gly	Tyr	Ala	Ser	Ile 230	Ser	Lys	Glu	Asn	Ile 235	Ser	Lys	Tyr	Ala	Gly 240
	Asp	Tyr	Ile	Phe	Leu 245	Ser	Lys	Pro	Ser	Tyr 250	Gly	Lys	Phe	Asp	Phe 255	Glu

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Lys Thr His Thr Trp Gln Asn Ile Glu Ala Val Lys Lys Gly His Val
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 5 Ile Ser Tyr Lys Ala Glu Asp Tyr Trp Phe Thr Asp Pro Ile Thr Leu
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 Lys Asp Asn Leu Asn Gly Glu Lys Pro Thr Thr Asn Leu Asn His Asn
 35 40 45
 Ile Thr Ser Pro Ser Val Asn Ser Glu Met Asn Asn Asn Glu Thr Gly
 50 55 60
 25 Thr Pro His Glu Ser Asn Gln Thr Gly Asn Glu Gly Thr Gly Ser Asn
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 Ser Arg Asp Ala Asn Pro Asp Ser Asn Asn Val Lys Pro Asp Ser Asn
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 210 215 220
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 245 250 255
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 260 265 270
 Lys Asp Tyr Gly Glu Val Thr Asp Glu Asp Ile Tyr Asn Ile Ile Arg
 275 280 285

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 305 310 315 320
 Tyr Tyr Arg Asn Leu Asp Glu Gln Val Leu Ala Leu Ile Thr Gly Glu
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 Ile Gly Ser Met Pro Asp Leu Lys Lys Pro Glu Asp Lys Pro Asp Ser
 340 345 350
 Lys Gln Arg Ser Phe Glu Pro His Glu Lys Asp Asp Phe Thr Val Val
 355 360 365
 Lys Lys Gln Glu Asp Asn Lys Lys Ser Ala Ser Thr Ala Tyr Ser Lys
 370 375 380
 Ser Trp Leu Ala Ile Val Cys Ser Met Met Val Val Phe Ser Ile Met
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 Asn Gln Lys Pro Ile Glu Ala Gly Ala Tyr Asn Tyr Thr Phe Asp Tyr
 50 55 60
 Glu Gly Phe Thr Tyr His Phe Glu Ser Asp Gly Thr His Phe Ala Trp
 65 70 75 80
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 85 90 95
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 Gln Ser Gln Glu Val Glu Ala Pro Gln Asn Ala Gln Thr Gln Gln Pro
 115 120 125
 Gln Ala Ser Thr Ser Asn Asn Ser Gln Val Thr Ala Thr Pro Thr Glu
 130 135 140
 Ser Lys Ser Ser Glu Gly Ser Ser Val Asn Val Asn Ala His Leu Lys
 145 150 155 160
 Gln Ile Ala Gln Arg Glu Ser Gly Gly Asn Ile His Ala Val Asn Pro
 165 170 175
 Thr Ser Gly Ala Ala Gly Lys Tyr Gln Phe Leu Gln Ser Thr Trp Asp
 180 185 190
 Ser Val Ala Pro Ala Lys Tyr Lys Gly Val Ser Pro Ala Asn Ala Pro

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	1				5					10					15			
15	Leu	Ala	Val	Ala	Ile	Ile	Ile	Gly	Ser	Ile	Ile	Met	Val	Val	Phe	Ser		
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	Gln	Arg	Gln	Ala	Gln	Thr	Phe	Lys	Ile	Gln	Gln	Gln	Gln	Phe	Val	Lys		
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20	Lys	Pro	Ile	Pro	Thr	Leu	Phe	Leu	His	Gly	Phe	Gly	Gly	Ser	Ala	Asn		
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	Ser	Glu	Lys	Phe	Met	Val	Lys	Gln	Ala	Glu	Lys	Arg	Gly	Val	Thr	Lys		
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25	Asp	Ile	Ile	Thr	Ala	Tyr	Val	Ser	Lys	Asp	Gly	Ala	Val	Thr	Phe	Lys		
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	Gly	Lys	Leu	Arg	Lys	Asp	Ala	Val	Asn	Pro	Ile	Val	Lys	Ile	Glu	Leu		
			100						105					110				
30	Glu	Asn	Asn	Arg	Gln	Gly	Tyr	Leu	Asp	Lys	Asn	Ala	Ala	Trp	Phe	Lys		
			115					120					125					
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45	Tyr	Gln	Gln	Leu	Arg	Val	Leu	Lys	Asp	Ile	Tyr	Lys	Gly	Lys	Gly	Ile		
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	Gln	His	Ser	Gln	Leu	His	Glu	Asn	Glu	Asn	Val	Ala	Asn	Glu	Leu	Ile		
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 Asp Lys His Gln Ile Ala Val Ala Asp Thr Asn Val Gln Thr Pro Asp
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 Lys Tyr Asp Glu Asn Asn Pro Asp Met Lys Lys Lys Phe Asp Ala Thr
 65 70 75 80

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 Glu Lys Glu Ala Thr Asn Leu Leu Lys Glu Met Lys Thr Glu Ser Gly
 85 90 95

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 Arg Lys Tyr Leu Trp Ser Gly Ala Glu Thr Leu Glu Thr Asn Ser Ser
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 Glu Lys Lys Lys Phe Thr Ala Pro Ile Lys Thr Phe Ala Pro Asp Ser
 210 215 220

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 275 280 285

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 Tyr Lys Asp Gly Ser Tyr Ile Asp His Gln Asp Val Pro Tyr Thr Gly
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105
 Ile Lys Glu Thr Pro Phe Asn Asp Lys Thr Gln Asn Asp Thr Thr Leu
 325 330 335

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 5 Met Met Asp Leu Ser Arg Gly Arg Ala Ile Ser Arg Glu Asn Glu Thr
 355 360 365
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 370 375 380
 10 Asp Ala Met Asp Asp Ser Thr Lys Ala Lys Tyr Lys Lys Ile Val Lys
 385 390 400
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 20 Asp Arg Val Thr Tyr His Asn Lys Asp Leu Asp Phe Ala Phe Gly Leu
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 Ser Met Thr Ser Lys Asn Val Ala Arg Tyr Glu Ser Ile Asn Gly Glu
 465 470 475 480
 Asn Leu Lys Gly Trp His Thr Gly Ala Gly Met Ser Tyr Leu Tyr Asn
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 565 570 575
 Phe Leu Gly Thr Gly Ile Lys Ser Thr Asp Ser Ser Lys Asn Pro Val
 580 585 590
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 Asp Lys Gln Thr Thr Asn Ser Asp Asn Gln Glu Asn Asn Ser Val Phe
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 Glu Val Thr Gln Lys His Ser Asn Ser Asp Asn Lys Tyr Gly Tyr Val
 675 680 685
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 740 745 750
 Leu Lys Lys Lys Asp Asp Asn Thr Tyr Glu Cys Ser Phe Tyr Asn Pro
 755 760 765
 10 Glu Ser Thr Asn Ser Ala Ser Asp Ile Glu Ser Lys Ile Ser Met Thr
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 50 55 60
 Thr Ala Gly Gln Cys Thr Trp Tyr Val Tyr Asp Lys Val Gly Gly Glu
 65 70 75 80
 35 Ile Gly Ser Thr Trp Gly Asn Ala Asn Asn Trp Ala Ala Ala Ala Gln
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 Gly Ala Gly Phe Thr Val Asn His Thr Pro Ser Lys Gly Ala Ile Leu
 100 105
 40 Gln Ser Ser Glu Gly Pro Phe Gly His Val Ala Tyr Val Glu Ser Val
 115 120 125
 Asn Ser Asp Gly Ser Val Thr Ile Ser Glu Met Asn Tyr Ser Gly Gly
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 Val Thr Ala Gln Ser Ile Gly Asp Gln Gln Thr Arg Glu Asn Ala Asn
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 5 Gln Leu Gly Arg Ser Ile Gly Ser Leu Trp Gly Asn Ala Asn Asn Trp
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	Ala	Ala	Phe	Asn	Lys	Ala	Ala	Gln	Met	Tyr	Gly	Ile	Asn	Glu	Val	Tyr	
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35	Leu	Ile	Ser	His	Ala	Leu	Leu	Glu	Thr	Gly	Asn	Gly	Thr	Ser	Gln	Leu	
					1125					1130					1135		
	Ala	Lys	Gly	Ala	Asp	Val	Val	Asn	Asn	Lys	Val	Val	Thr	Asn	Ser	Asn	
				1140					1145					1150			
40	Thr	Lys	Tyr	His	Asn	Val	Phe	Gly	Ile	Ala	Ala	Tyr	Asp	Asn	Asp	Pro	
			1155					1160					1165				
	Leu	Arg	Glu	Gly	Ile	Lys	Tyr	Ala	Lys	Gln	Ala	Gly	Trp	Asp	Thr	Val	
		1170					1175					1180					
45	Ser	Lys	Ala	Ile	Val	Gly	Gly	Ala	Lys	Phe	Ile	Gly	Asn	Ser	Tyr	Val	
		1185					1190					1195				1200	
	Lys	Ala	Gly	Gln	Asn	Thr	Leu	Tyr	Lys	Met	Arg	Trp	Asn	Pro	Ala	His	
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50	Pro	Gly	Thr	His	Gln	Tyr	Ala	Thr	Asp	Val	Asp	Trp	Ala	Asn	Ile	Asn	
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	Ala	Lys	Ile	Ile	Lys	Gly	Tyr	Tyr	Asp	Lys	Ile	Gly	Glu	Val	Gly	Lys	
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10	Asn	Lys	Ile	Asn	His	Glu	Ala	His	Ile	Val	Gly	Val	Lys	Asn	Asp	Lys
			35					40					45			
	Asn	Glu	Val	Ile	Ala	Ala	Cys	Leu	Leu	Thr	Glu	Ala	Arg	Ile	Phe	Lys
		50					55					60				
15	Phe	Tyr	Lys	Tyr	Phe	Tyr	Ser	His	Arg	Gly	Pro	Leu	Leu	Asp	Tyr	Phe
	65					70					75					80
	Asp	Ala	Lys	Leu	Val	Cys	Tyr	Phe	Phe	Lys	Glu	Leu	Ser	Lys	Phe	Ile
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20	Tyr	Lys	Asn	Arg	Gly	Val	Phe	Ile	Leu	Val	Asp	Pro	Tyr	Leu	Ile	Glu
				100					105					110		
	Asn	Leu	Arg	Asp	Ala	Asn	Gly	Arg	Ile	Ile	Lys	Asn	Tyr	Asn	Asn	Ser
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25	Val	Ile	Val	Lys	Met	Leu	Gly	Lys	Ile	Gly	Tyr	Leu	His	Gln	Gly	Tyr
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	Thr	Thr	Gly	Tyr	Ser	Asn	Lys	Ser	Gln	Ile	Arg	Trp	Ile	Ser	Val	Leu
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30	Asp	Leu	Lys	Asp	Lys	Asp	Glu	Asn	Gln	Leu	Leu	Lys	Glu	Met	Glu	Tyr
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	Gln	Thr	Arg	Arg	Asn	Ile	Lys	Lys	Thr	Ile	Glu	Ile	Gly	Val	Lys	Val
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35	Glu	Asp	Leu	Ser	Ile	Glu	Glu	Thr	Asn	Arg	Phe	Tyr	Lys	Leu	Phe	Gln
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	Met	Ala	Glu	Glu	Lys	His	Gly	Phe	His	Phe	Met	Asn	Glu	Asp	Tyr	Phe
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	Cys	Ile	Asn	Leu	Asn	Glu	Tyr	Gln	Asp	Lys	Leu	Lys	Ile	Gln	Leu	Leu
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45	Lys	Ile	Glu	Asn	Glu	Met	Met	Thr	Val	Asn	Arg	Ala	Leu	Asn	Glu	Asn
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	Pro	Asn	Ser	Lys	Lys	Asn	Lys	Ser	Lys	Leu	Asn	Gln	Leu	Asn	Met	Gln
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	Asp	Asp	Glu	Val	Tyr	Tyr	Leu	Ser	Ser	Gly	Ser	Asn	Pro	Lys	Tyr	Asn
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Lys Ser His Asn Ile Asn Arg Tyr Asn Phe Tyr Gly Ile Thr Gly Val
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 Phe Ser Asn Glu Ala Asp Asp Phe Gly Val Gln Gln Phe Lys Lys Gly
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 35 40 45
 Gln Phe Lys Pro Leu Gly Gln Asp Gln Phe Val Thr Val Pro Asp Gly
 50 55 60
 Val Ala His Phe Leu Glu His Lys Leu Phe Glu Lys Glu Glu Glu Asp
 25 65 70 75 80
 Leu Phe Thr Ala Phe Ala Glu Asp Asn Ala Gln Ala Asn Ala Phe Thr
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 30 Ser Phe Asp Arg Thr Ser Tyr Leu Phe Ser Ala Thr Asp Asn Ile Glu
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 35 Lys Glu Thr Val Asp Lys Glu Lys Gly Ile Ile Ala Glu Glu Ile Lys
 130 135 140
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 40 Ala Met Tyr Gln Gln His Pro Ile Arg Val Asp Ile Ala Gly Ser Val
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 45 Thr Phe Tyr His Pro Ser Asn Met Val Leu Phe Val Val Gly Asp Val
 195 200 205
 Asp Pro Glu Ala Ile Cys Arg Ile Val Lys Gln His Glu Asp Ala Arg
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 50 Asn Lys Val Asn Gln Pro Lys Ile Glu Arg Gly Leu Val Asp Glu Pro
 225 230 235
 Glu Asp Val Lys Glu Ala Phe Val Thr Glu Ser Met Lys Ile Gln Ser
 245 250 255
 55 Pro Arg Leu Met Leu Gly Phe Lys Asn Lys Pro Leu Gln Glu Ala Pro
 260 265 270
 Gln Lys Tyr Val Gln Arg Asp Leu Glu Met Ser Leu Phe Phe Glu Leu

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	Ser	Phe	Ser	Ile	Val	Thr	Ser	Ala	Thr	Glu	Glu	Pro	Asp	Lys	Leu	Lys
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	Ala	Glu	Ala	Phe	Glu	Leu	Leu	Lys	Lys	Gln	Phe	Ile	Gly	Glu	Phe	Ile
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	Tyr	Phe	Glu	Gly	Val	Ser	Val	Phe	Asp	Met	Leu	Asp	Ile	Val	Glu	Asn
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20	Ile	Thr	Leu	Asp	Ser	Ile	Asn	Glu	Thr	Ser	Ser	Leu	Tyr	Leu	Asn	Leu
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35	Ser	Gln	Ala	Arg	Gln	Thr	Ala	Glu	Asp	Ile	Val	Asn	Gln	Ala	His	Lys
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	Ser	Glu	Leu	Gln	Arg	Gln	Glu	Thr	Arg	Leu	Leu	Gln	Lys	Glu	Glu	Asn
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45	Leu	Glu	Arg	Lys	Ser	Asp	Leu	Leu	Asp	Lys	Lys	Asp	Glu	Ile	Leu	Glu
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	Gln	Lys	Glu	Ser	Lys	Ile	Glu	Glu	Lys	Gln	Gln	Gln	Val	Asp	Ala	Lys
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55	Arg	Ile	Ser	Gly	Leu	Thr	Gln	Glu	Glu	Ala	Ile	Asn	Glu	Gln	Leu	Gln
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	Arg	Val	Glu	Glu	Glu	Leu	Ser	Gln	Asp	Ile	Ala	Val	Leu	Val	Lys	Glu
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55	Lys	Glu	Lys	Glu	Ala	Lys	Glu	Lys	Val	Asp	Lys	Thr	Ala	Lys	Glu	Leu
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Leu Ala Thr Ala Val Gln Arg Leu Ala Ala Asp His Thr Ser Glu Ser
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 5 Thr Val Ser Val Val Asn Leu Pro Asn Asp Glu Met Lys Gly Arg Ile
 210 215 220
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 225 230 235 240
 10 Ile Asp Leu Ile Ile Asp Asp Thr Pro Glu Ala Val Ile Leu Ser Gly
 245 250 255
 Phe Asp Pro Ile Arg Arg Glu Ile Ala Arg Thr Ala Leu Val Asn Leu
 260 270
 15 Val Ser Asp Gly Arg Ile His Pro Gly Arg Ile Glu Asp Met Val Glu
 275 280 285
 Lys Ala Arg Lys Glu Val Asp Asp Ile Ile Arg Glu Ala Gly Glu Gln
 290 295 300
 20 Ala Thr Phe Glu Val Asn Ala His Asn Met His Pro Asp Leu Val Lys
 305 310 315 320
 Ile Val Gly Arg Leu Asn Tyr Arg Thr Ser Tyr Gly Gln Asn Val Leu
 325 330 335
 25 Lys His Ser Ile Glu Val Ala His Leu Ala Ser Met Leu Ala Ala Glu
 340 345 350
 Leu Gly Glu Asp Glu Thr Leu Ala Lys Arg Ala Gly Leu Leu His Asp
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 30 Val Gly Lys Ala Ile Asp His Glu Val Glu Gly Ser His Val Glu Ile
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 Gly Val Glu Leu Ala Lys Lys Tyr Gly Glu Asn Glu Thr Val Ile Asn
 385 390 395 400
 35 Ala Ile His Ser His His Gly Asp Val Glu Pro Thr Ser Ile Ile Ser
 405 410 415
 Ile Leu Val Ala Ala Ala Asp Ala Leu Ser Ala Ala Arg Pro Gly Ala
 420 425 430
 40 Arg Lys Glu Thr Leu Glu Asn Tyr Ile Arg Arg Leu Glu Arg Leu Glu
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 Thr Leu Ser Glu Ser Tyr Asp Gly Val Glu Lys Ala Phe Ala Ile Gln
 450 455 460
 45 Ala Gly Arg Glu Ile Arg Val Ile Val Ser Pro Glu Glu Ile Asp Asp
 465 470 475 480
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 5 20 30
 Val Tyr Ala Arg Lys Gln Leu Ile Lys Lys Asn Lys Ala Leu Ser Ala
 35 40 45
 Glu Asn Ala Glu Leu Arg Ser Gln Met Leu Ser Ser Asn Asn Asp Val
 10 50 55 60
 Gly His His Ala Tyr Lys Asn Ala Lys Arg Glu Leu Arg Lys Ile Leu
 65 70 75 80
 Asp Ser Tyr Leu Glu Asn Gly Lys Leu Lys Tyr Tyr Asp Ile Ile Val
 15 85 90 95
 Thr Ser Asn Leu Ala Thr Lys His Pro Phe Phe Glu Tyr Ala Arg Ser
 100 105 110
 Phe Asp Phe Ile Ile Val Ser Asp Ile Gly Leu Ile Asn Val Asp Val
 115 120 125
 Lys Ser Trp Gly Glu Lys Thr Phe Tyr His Phe Asp Val Pro Asp Glu
 130 135 140
 His Asp Thr Glu Ile Ser Asn Ser Asn Ile Glu Lys Val Val Gly His
 145 150 155 160
 Tyr Ile Ser Gln Gln Tyr His Asp Gln Phe Asn Ser Ser Arg Lys Ser
 165 170 175
 Ile Tyr Thr Phe Thr Glu Thr Val Gln Pro Asn Arg Val Ile Tyr Asp
 180 185 190
 Phe Tyr Asp Tyr Asp Pro Tyr Gln Leu Ala Ala Asn Asn Ala Lys Ala
 195 200 205
 Leu Lys Asp His Ile Glu Gln Asn Phe Asn Phe Lys Val Gln Ser Thr
 210 215 220
 Gly Val Ile Tyr Phe Ser Asp Gly Thr Val Asn Ile Ile Gln Gly Ser
 225 230 235 240
 Glu Glu Arg Asp Lys Tyr Val Asp Thr Val Ser Thr Lys Ser Ser Leu
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 Arg Pro Arg Gly Lys Phe Arg Phe Pro Met Ser Val Ala Tyr Glu Asn

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	Glu 65	Asn	Val	Glu	Gln	Ser 70	Ala	Asp	Thr	Ile	Ser 75	Asp	Glu	Lys	Glu	Gln 80	
5	Tyr	His	Arg	Asp	Tyr 85	Arg	Lys	Gln	Ser	His 90	Asp	Ser	Arg	Ser	Gln 95	Lys	
	Arg	His	Arg	Arg 100	Arg	Arg	Asn	Gln	Thr 105	Thr	Glu	Glu	Gln	Asn 110	Tyr	Ser	
10	Glu	Gln	Arg 115	Gly	Asn	Ser	Lys	Ile 120	Ser	Gln	Gln	Ser	Ile 125	Lys	Tyr	Lys	
	Asp	His 130	Ser	His	Tyr	His	Thr 135	Asn	Lys	Pro	Gly	Thr 140	Tyr	Val	Ser	Ala	
15	Ile 145	Asn	Gly	Ile	Glu	Lys 150	Glu	Thr	His	Lys	Pro 155	Lys	Thr	His	Asn	Met 160	
	Tyr	Ser	Asn	Asn	Thr 165	Asn	His	Arg	Ala	Lys 170	Asp	Ser	Thr	Pro	Asp 175	Tyr	
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25	Pro	Ser 210	Glu	Lys	Val	Glu	Ser 215	Asp	Lys	Gln	Lys	Tyr 220	Asp	Lys	Tyr	Val	
	Ala 225	Lys	Thr	Gln	Thr	Ser 230	Gln	Asn	Lys	Gln	Leu 235	Glu	Gln	Glu	Lys	Gln 240	
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	Ile	Arg 290	Arg	Glu	Arg	Glu	Arg 295	Lys	Val	Leu	Gln	Lys 300	Arg	Arg	Phe	Lys	
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45	Tyr	Val	Gly	Asp 340	Ser	Ser	Leu	Asn	Asp 345	Asp	Ser	Asp	Leu	Thr 350	Asp	Asn	
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50	Val	Ser 370	Asn	Asp	Glu	Asn	Lys 375	Gln	Ala	Ser	Ile	Gln 380	Asn	Glu	Asp	Thr	
	Asn 385	Asp	Thr	His	Val	Asp 390	Glu	Ser	Pro	Tyr	Asn 395	Tyr	Glu	Glu	Val	Ser 400	
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5
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Glu Val Asn Asp Lys Asp Glu Leu Lys Asn Gln Ser Arg Leu Ile Ala
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 Gln Ile Asp Asp Ala Glu Phe Tyr Glu Leu Asn Asp Thr Glu Val Asp
 465 470 475 480
 Glu Asp Thr Thr Ser Asn Ile Glu Asp Asn Thr Asn Arg Asn Ala Ser
 485 490 495
 Glu Met His Val Asp Ala Pro Lys Thr Gln Glu Tyr Ala Val Thr Glu
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 Ser Gln Val Asn Asn Ile Asp Lys Thr Val Asp Asn Glu Ile Glu Leu
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 Ala Pro Arg His Lys Lys Asp Asp Gln Thr Asn Leu Ser Val Asn Ser
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 Leu Lys Thr Asn Asp Val Asn Asp Asn His Val Val Glu Asp Ser Ser
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 Met Asn Glu Ile Glu Lys Asn Asn Ala Glu Ile Thr Glu Asn Val Gln
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 595 600 605
 Arg Pro Phe Asn Val Val Met Thr Pro Ser Asp Lys Lys Arg Met Met
 610 615 620
 Asp Arg Lys Lys His Ser Lys Val Asn Val Pro Glu Leu Lys Pro Val
 625 630 635 640
 Gln Ser Lys Gln Ala Val Ser Glu Arg Met Pro Ala Ser Gln Ala Thr
 645 650 655
 Pro Ser Ser Arg Ser Asp Ser Gln Glu Ser Asn Thr Asn Ala Tyr Lys
 660 665 670
 Thr Asn Asn Met Thr Ser Asn Asn Val Glu Asn Asn Gln Leu Ile Gly
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 His Ala Glu Thr Glu Asn Asp Tyr Gln Asn Ala Gln Gln Tyr Ser Glu
 690 695 700
 Gln Lys Pro Ser Val Asp Ser Thr Gln Thr Glu Ile Phe Glu Glu Ser
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 Gln Asp Asp Asn Gln Leu Glu Asn Glu Gln Val Asp Gln Ser Thr Ser
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 740 745 750
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 Lys Asp Leu Gln Ser Ser Phe Ser Asn Lys Asn Glu Asp Thr Ala Asn
 770 775 780
 Glu Asn Arg Pro Arg Thr Asn Gln Gln Asp Val Ala Thr Asn Gln Ala
 785 790 795 800

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Val Gln Thr Ser Lys₈₀₅ Pro Met Ile Arg Lys₈₁₀ Gly Pro Asn Ile Lys₈₁₅ Leu
 Pro Ser Val Ser₈₂₀ Leu Leu Glu Glu Pro₈₂₅ Gln Val Ile Glu Ser₈₃₀ Asp Glu
 5 Asp Trp Ile₈₃₅ Thr Asp Lys Lys Lys₈₄₀ Glu Leu Asn Asp Ala₈₄₅ Leu Phe Tyr
 Phe Asn Val₈₅₀ Pro Ala Glu Val₈₅₅ Gln Asp Val Thr Glu₈₆₀ Gly Pro Ser Val
 10 Thr Arg Phe Glu Leu Ser₈₇₀ Val Glu Lys Gly Val₈₇₅ Lys Val Ser Arg Ile₈₈₀
 Thr Ala Leu Gln Asp₈₈₅ Asp Ile Lys Met Ala₈₉₀ Leu Ala Ala Lys Asp₈₉₅ Ile
 15 Arg Ile Glu Ala₉₀₀ Pro Ile Pro Gly Thr Ser Arg Val Gly Ile₉₁₀ Glu Val
 Pro Asn Gln₉₁₅ Asn Pro Thr Thr Val₉₂₀ Asn Leu Arg Ser Ile₉₂₅ Ile Glu Ser
 20 Pro Ser Phe Lys Asn Ala Glu₉₃₅ Ser Lys Leu Thr Val₉₄₀ Ala Met Gly Tyr
 Arg Ile Asn Asn Glu Pro Leu Leu Met Asp Ile₉₅₅ Ala Lys Thr Pro His₉₆₀
 25 Ala Leu Ile Ala Gly₉₆₅ Ala Thr Gly Ser Gly₉₇₀ Lys Ser Val Cys Ile₉₇₅ Asn
 Ser Ile Leu Met₉₈₀ Ser Leu Leu Tyr Lys₉₈₅ Asn His Pro Glu Glu₉₉₀ Leu Arg
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 Leu Pro His Leu Val Ala Pro Val Ile Thr Asp Val Lys Ala Ala Thr
 35 Gln Ser Leu Lys Trp Ala Val Glu Glu Met Glu Arg Arg Tyr Lys Leu₁₀₄₀
 Phe Ala His Tyr His Val Arg Asn Ile Thr Ala Phe Asn Lys Lys Ala₁₀₅₅
 40 Pro Tyr Asp Glu Arg Met Pro Lys Ile Val Ile Val Ile Asp Glu Leu
 Ala Asp Leu Met Met Met Ala Pro Gln Glu Val Glu Gln Ser Ile Ala
 45 Arg Ile Ala Gln Lys Ala Arg Ala Cys Gly Ile His Met Leu Val Ala
 Thr Gln Arg Pro Ser Val Asn Val Ile Thr Gly Leu Ile Lys Ala Asn₁₁₂₀
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 Thr Ile Leu Asp Ser Gly Gly Ala Glu Arg Leu Leu Gly Tyr Gly Asp
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 55 Thr Phe Val Ser Asp Asp Glu Ile Asp Asp Val Val Asp Phe Ile Lys

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				20					25					30					
	Thr	Ala	Lys	Ala	Glu	Asn	Thr	Val	Lys	Gln	Ile	Thr	Asn	Thr	Asn	Val			
			35					40					45						
30	Ala	Pro	Tyr	Ser	Gly	Val	Thr	Trp	Met	Gly	Ala	Gly	Thr	Gly	Phe	Val			
		50					55					60							
	Val	Gly	Asn	His	Thr	Ile	Ile	Thr	Asn	Lys	His	Val	Thr	Tyr	His	Met			
	65					70					75				80				
35	Lys	Val	Gly	Asp	Glu	Ile	Lys	Ala	His	Pro	Asn	Gly	Phe	Tyr	Asn	Asn			
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	Gly	Gly	Gly	Leu	Tyr	Lys	Val	Thr	Lys	Ile	Val	Asp	Tyr	Pro	Gly	Lys			
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40	Glu	Asp	Ile	Ala	Val	Val	Gln	Val	Glu	Glu	Lys	Ser	Thr	Gln	Pro	Lys			
			115				120						125						
	Gly	Arg	Lys	Phe	Lys	Asp	Phe	Thr	Ser	Lys	Phe	Asn	Ile	Ala	Ser	Glu			
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45	Ala	Lys	Glu	Asn	Glu	Pro	Ile	Ser	Val	Ile	Gly	Tyr	Pro	Asn	Pro	Asn			
	145					150					155					160			
	Gly	Asn	Lys	Leu	Gln	Met	Tyr	Glu	Ser	Thr	Gly	Lys	Val	Leu	Ser	Val			
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50	Asn	Gly	Asn	Ile	Val	Ser	Ser	Asp	Ala	Ile	Ile	Gln	Pro	Gly	Ser	Ser			
				180					185					190					
	Gly	Ser	Pro	Ile	Leu	Asn	Ser	Lys	His	Glu	Ala	Ile	Gly	Val	Ile	Tyr			
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55	Ala	Gly	Asn	Lys	Pro	Ser	Gly	Glu	Ser	Thr	Arg	Gly	Phe	Ala	Val	Tyr			
		210					215					220							
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 <211> 238
 <212> PRT
 <213> Staphylococcus aureus

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 Thr Ala Lys Ala Glu His Asn Val Lys Leu Ile Lys Asn Thr Asn Val
 35 40 45
 Ala Pro Tyr Asn Gly Val Val Ser Ile Gly Ser Gly Thr Gly Phe Ile
 15 50 55 60
 Val Gly Lys Asn Thr Ile Val Thr Asn Lys His Val Val Ala Gly Met
 65 70 75 80
 Glu Ile Gly Ala His Ile Ile Ala His Pro Asn Gly Glu Tyr Asn Asn
 20 85 90 95
 Gly Gly Phe Tyr Lys Val Lys Lys Ile Val Arg Tyr Ser Gly Gln Glu
 100 105
 Asp Ile Ala Ile Leu His Val Glu Asp Lys Ala Val His Pro Lys Asn
 115 120 125
 25 Arg Asn Phe Lys Asp Tyr Thr Gly Ile Leu Lys Ile Ala Ser Glu Ala
 130 135 140
 Lys Glu Asn Glu Arg Ile Ser Ile Val Gly Tyr Pro Glu Pro Tyr Ile
 145 150 155 160
 30 Asn Lys Phe Gln Met Tyr Glu Ser Thr Gly Lys Val Leu Ser Val Lys
 165 170 175
 Gly Asn Met Ile Ile Thr Asp Ala Phe Val Glu Pro Gly Asn Ser Gly
 180 185 190
 35 Ser Ala Val Phe Asn Ser Lys Tyr Glu Val Val Gly Val His Phe Gly
 195 200 205
 Gly Asn Gly Pro Gly Asn Lys Ser Thr Lys Gly Tyr Gly Val Tyr Phe
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 40 Ser Pro Glu Ile Lys Lys Phe Ile Ala Asp Asn Thr Asp Lys
 225 230 235

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 <213> Staphylococcus aureus

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 20 25 30
 Ile Ala Asn Ala Glu Lys Asn Val Thr Gln Val Lys Asp Thr Asn Ile
 35 40 45
 55 Phe Pro Tyr Asn Gly Val Val Ser Phe Lys Asp Ala Thr Gly Phe Val
 50 55 60

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Ile Gly Lys Asn Thr Ile Ile Thr Asn Lys His Val Ser Lys Asp Tyr
 65 70 75 80
 Lys Val Gly Asp Arg Ile Thr Ala His Pro Asn Gly Asp Lys Gly Asn
 5 85 90 95
 Gly Gly Ile Tyr Lys Ile Lys Ser Ile Ser Asp Tyr Pro Gly Asp Glu
 100 105 110
 Asp Ile Ser Val Met Asn Ile Glu Glu Gln Ala Val Glu Arg Gly Pro
 115 120 125
 Lys Gly Phe Asn Phe Asn Glu Asn Val Gln Ala Phe Asn Phe Ala Lys
 130 135 140
 Asp Ala Lys Val Asp Asp Lys Ile Lys Val Ile Gly Tyr Pro Leu Pro
 145 150 155 160
 Ala Gln Asn Ser Phe Lys Gln Phe Glu Ser Thr Gly Thr Ile Lys Arg
 165 170 175
 Ile Lys Asp Asn Ile Leu Asn Phe Asp Ala Tyr Ile Glu Pro Gly Asn
 180 185 190
 Ser Gly Ser Pro Val Leu Asn Ser Asn Asn Glu Val Ile Gly Val Val
 195 200 205
 Tyr Gly Gly Ile Gly Lys Ile Gly Ser Glu Tyr Asn Gly Ala Val Tyr
 210 215 220
 Phe Thr Pro Gln Ile Lys Asp Phe Ile Gln Lys His Ile Glu Gln
 225 230 235
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 <212> PRT
 <213> Staphylococcus aureus
 30
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 Thr Ala Lys Ala Glu Asn Asn Val Thr Lys Val Lys Asp Thr Asn Ile
 35 35 40 45
 Phe Pro Tyr Thr Gly Val Val Ala Phe Lys Ser Ala Thr Gly Phe Val
 50 50 55 60
 Val Gly Lys Asn Thr Ile Leu Thr Asn Lys His Val Ser Lys Asn Tyr
 65 70 75 80
 Lys Val Gly Asp Arg Ile Thr Ala His Pro Asn Ser Asp Lys Gly Asn
 85 90 95
 Gly Gly Ile Tyr Ser Ile Lys Lys Ile Ile Asn Tyr Pro Gly Lys Glu
 100 105 110
 Asp Val Ser Val Ile Gln Val Glu Glu Arg Ala Ile Glu Arg Gly Pro
 115 120 125
 Lys Gly Phe Asn Phe Asn Asp Asn Val Thr Pro Phe Lys Tyr Ala Ala
 130 135 140
 Gly Ala Lys Ala Gly Glu Arg Ile Lys Val Ile Gly Tyr Pro His Pro
 145 150 155 160
 Tyr Lys Asn Lys Tyr Val Leu Tyr Glu Ser Thr Gly Pro Val Met Ser

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					165					170					175	
	Val	Glu	Gly	Ser	Ser	Ile	Val	Tyr	Ser	Ala	His	Thr	Glu	Ser	Gly	Asn
				180					185					190		
5	Ser	Gly	Ser	Pro	Val	Leu	Asn	Ser	Asn	Asn	Glu	Leu	Val	Gly	Ile	His
			195					200					205			
	Phe	Ala	Ser	Asp	Val	Lys	Asn	Asp	Asp	Asn	Arg	Asn	Ala	Tyr	Gly	Val
		210					215					220				
10	Tyr	Phe	Thr	Pro	Glu	Ile	Lys	Lys	Phe	Ile	Ala	Glu	Asn	Ile	Asp	Lys
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20	Thr	Ser	Leu	Gly	Phe	Ala	Glu	Asn	Ile	Ser	Asn	Gln	Pro	His	Ser	Ile
			20						25					30		
	Ala	Lys	Ala	Glu	Lys	Asn	Val	Lys	Glu	Ile	Thr	Asp	Ala	Thr	Lys	Glu
			35					40					45			
25	Pro	Tyr	Asn	Ser	Val	Val	Ala	Phe	Val	Gly	Gly	Thr	Gly	Val	Val	Val
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	Gly	Lys	Asn	Thr	Ile	Val	Thr	Asn	Lys	His	Ile	Ala	Lys	Ser	Asn	Asp
	65				70						75					80
30	Ile	Phe	Lys	Asn	Arg	Val	Ser	Ala	His	His	Ser	Ser	Lys	Gly	Lys	Gly
					85					90					95	
	Gly	Gly	Asn	Tyr	Asp	Val	Lys	Asp	Ile	Val	Glu	Tyr	Pro	Gly	Lys	Glu
				100					105					110		
35	Asp	Leu	Ala	Ile	Val	His	Val	His	Glu	Thr	Ser	Thr	Glu	Gly	Leu	Asn
			115					120					125			
	Phe	Asn	Lys	Asn	Val	Ser	Tyr	Thr	Lys	Phe	Ala	Asp	Gly	Ala	Lys	Val
		130					135					140				
40	Lys	Asp	Arg	Ile	Ser	Val	Ile	Gly	Tyr	Pro	Lys	Gly	Ala	Gln	Thr	Lys
	145					150					155					160
	Tyr	Lys	Met	Phe	Glu	Ser	Thr	Gly	Thr	Ile	Asn	His	Ile	Ser	Gly	Thr
				165						170					175	
45	Phe	Met	Glu	Phe	Asp	Ala	Tyr	Ala	Gln	Pro	Gly	Asn	Ser	Gly	Ser	Pro
				180					185					190		
	Val	Leu	Asn	Ser	Lys	His	Glu	Leu	Ile	Gly	Ile	Leu	Tyr	Ala	Gly	Ser
			195					200					205			
50	Gly	Lys	Asp	Glu	Ser	Glu	Lys	Asn	Phe	Gly	Val	Tyr	Phe	Thr	Pro	Gln
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	Leu	Lys	Glu	Phe	Ile	Gln	Asn	Asn	Ile	Glu	Lys					
	225					230					235					
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	<211>	163														
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55	<213>	Staphylococcus aureus														

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 Ser Phe Ser Ser Ile Thr Asn Glu Val Ser Ala Ser Ser Ser Phe Asp
 5 20 25 30
 Lys Gly Lys Tyr Lys Lys Gly Asp Asp Ala Ser Tyr Phe Glu Pro Thr
 35 40 45
 Gly Pro Tyr Leu Met Val Asn Val Thr Gly Val Asp Gly Lys Gly Asn
 10 50 55 60
 Glu Leu Leu Ser Pro His Tyr Val Glu Phe Pro Ile Lys Pro Gly Thr
 65 70 75 80
 Thr Leu Thr Lys Glu Lys Ile Glu Tyr Tyr Val Glu Trp Ala Leu Asp
 15 85 90 95
 Ala Thr Ala Tyr Lys Glu Phe Arg Val Val Glu Leu Asp Pro Ser Ala
 100 105 110
 Lys Ile Glu Val Thr Tyr Tyr Asp Lys Asn Lys Lys Lys Glu Glu Thr
 20 115 120 125
 Lys Ser Phe Pro Ile Thr Glu Lys Gly Phe Val Val Pro Asp Leu Ser
 130 135 140
 Glu His Ile Lys Asn Pro Gly Phe Asn Leu Ile Thr Lys Val Ile Ile
 25 145 150 155 160
 Glu Lys Lys

 <210> 121
 <211> 290
 <212> PRT
 <213> Staphylococcus aureus

 <400> 121
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 Leu Ala Gly Cys Asp Tyr Ser Lys Pro Glu Lys Arg Ser Gly Phe Phe
 20 25 30
 Tyr Asn Thr Phe Val Asp Pro Met Lys Asn Val Leu Asp Trp Leu Gly
 35 40 45
 Asn Asn Leu Leu Asn Asp Asn Tyr Gly Leu Ala Ile Ile Ile Leu Val
 50 55 60
 Leu Val Ile Arg Ile Ile Leu Leu Pro Phe Met Leu Ser Asn Tyr Lys
 65 70 75 80
 Asn Ser His Met Met Arg Gln Lys Met Lys Val Ala Lys Pro Glu Val
 85 90 95
 Glu Lys Ile Gln Glu Lys Val Lys Arg Ala Arg Thr Gln Glu Glu Lys
 100 105 110
 Met Ala Ala Asn Gln Glu Leu Met Gln Val Tyr Lys Lys Tyr Asp Met
 115 120 125
 Asn Pro Ile Lys Ser Met Leu Gly Cys Leu Pro Met Leu Ile Gln Leu
 130 135 140
 Pro Ile Ile Met Gly Leu Tyr Phe Val Leu Lys Asp Gln Leu Val Asp
 145 150 155 160

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Gly Leu Phe Lys Tyr Pro His Phe Leu Trp Phe Asp Leu Gly Arg Pro
 165 170 175
 5 Asp Ile Trp Ile Thr Ile Ile Ala Gly Val Leu Tyr Phe Ile Gln Ala
 180 185 190
 Tyr Val Ser Ser Lys Thr Met Pro Asp Glu Gln Arg Gln Met Gly Tyr
 195 200 205
 10 Met Met Met Val Ile Ser Pro Ile Met Ile Ile Trp Ile Ser Leu Ser
 210 215 220
 Ser Ala Ser Ala Leu Gly Leu Tyr Trp Ser Val Ser Ala Ala Phe Leu
 225 230 235
 15 Val Val Gln Thr His Phe Ala Asn Ile Tyr Tyr Glu Lys Val Ala Lys
 245 250 255
 Lys Glu Val Gln Pro Phe Ile Glu Ala Tyr Glu Arg Glu His Asn Gly
 260 265 270
 20 Gly Ser Asn Lys Lys Gly Lys Asn Thr Gln Val Val Ser Lys Lys Lys
 275 280 285
 Lys Lys
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 <210> 122
 <211> 460
 <212> PRT
 <213> Staphylococcus aureus
 <400> 122
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 Arg Lys Ser Thr Asp Glu Asp Ile Gln Thr Asn Asn Ile Lys Met Arg
 35 40 45
 Lys Met Val Pro Trp Ala Ile Gly Phe Phe Ile Leu Ile Leu Ile Ile
 50 55 60
 Ile Leu Phe Phe Leu Leu Arg Asn Phe Asn Ser Pro Glu Ala Gln Thr
 65 70 75 80
 40 Lys Ile Leu Val Asn Ala Ile Glu Asn Asn Asp Lys Gln Lys Val Ala
 85 90 95
 Thr Leu Leu Ser Thr Lys Asp Asn Lys Val Asp Ser Glu Glu Ala Lys
 100 105 110
 45 Val Tyr Ile Asn Tyr Ile Lys Asp Glu Val Gly Leu Lys Gln Phe Val
 115 120 125
 Ser Asp Leu Lys Asn Thr Val His Lys Leu Asn Lys Ser Lys Thr Ser
 130 135 140
 50 Val Ala Ser Tyr Ile Gln Thr Arg Ser Gly Gln Asn Ile Leu Arg Val
 145 150 155 160
 Ser Lys Asn Gly Thr Arg Tyr Ile Phe Phe Asp Asn Met Ser Phe Thr
 165 170 175
 55 Ala Pro Thr Lys Gln Pro Ile Val Lys Pro Lys Glu Lys Thr Lys Tyr
 180 185 190
 Glu Phe Lys Ser Gly Gly Lys Lys Lys Met Val Ile Ala Glu Ala Asn

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		195					200				205					
	Lys	Val	Thr	Pro	Ile	Gly	Asn	Phe	Ile	Pro	Gly	Thr	Tyr	Arg	Ile	Pro
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5	Ala	Met	Lys	Ser	Thr	Glu	Asn	Gly	Asp	Phe	Ala	Gly	His	Leu	Lys	Phe
	225					230					235					240
	Asp	Phe	Arg	Gln	Ser	Asn	Ser	Glu	Thr	Val	Asp	Val	Thr	Glu	Asp	Phe
					245					250					255	
10	Glu	Glu	Ala	Asn	Ile	Ser	Val	Thr	Leu	Lys	Gly	Asp	Thr	Lys	Leu	Asn
				260					265					270		
	Asp	Ser	Ser	Lys	Lys	Val	Thr	Ile	Asn	Asp	His	Glu	Met	Ala	Phe	Ser
			275					280					285			
15	Ser	Ser	Lys	Thr	Tyr	Gly	Pro	Tyr	Pro	Gln	Asn	Lys	Asp	Ile	Thr	Ile
		290					295					300				
	Ser	Ala	Ser	Gly	Lys	Ala	Lys	Asp	Lys	Thr	Phe	Thr	Thr	Gln	Thr	Lys
	305					310					315					320
20	Thr	Ile	Lys	Ala	Ser	Asp	Leu	Lys	Tyr	Asn	Thr	Glu	Ile	Thr	Leu	Asn
					325					330					335	
	Phe	Asp	Ser	Glu	Asp	Ile	Glu	Asp	Tyr	Val	Glu	Lys	Lys	Glu	Lys	Glu
				340					345					350		
25	Glu	Asn	Ser	Leu	Lys	Asn	Lys	Leu	Ile	Glu	Phe	Phe	Ala	Gly	Tyr	Ser
			355					360					365			
	Leu	Ala	Asn	Asn	Ala	Ala	Phe	Asn	Gln	Ser	Asp	Phe	Asp	Phe	Val	Ser
	370						375					380				
30	Ser	Tyr	Ile	Lys	Lys	Gly	Ser	Ser	Phe	Tyr	Asp	Asp	Val	Lys	Lys	Arg
	385					390					395					400
	Val	Ser	Lys	Gly	Ser	Leu	Met	Met	Ile	Ser	Ser	Pro	Gln	Ile	Ile	Asp
					405					410					415	
35	Ala	Glu	Lys	His	Gly	Asp	Lys	Ile	Thr	Ala	Thr	Val	Arg	Leu	Ile	Asn
				420					425					430		
	Glu	Asn	Gly	Lys	Gln	Val	Asp	Lys	Glu	Tyr	Glu	Leu	Glu	Gln	Gly	Ser
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40	Gln	Asp	Arg	Leu	Gln	Leu	Ile	Lys	Thr	Ser	Glu	Lys				
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	<213>	Staphylococcus aureus														
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50	Tyr	Ala	His	Ile	Arg	Ile	Lys	Glu	Lys	Arg	Ser	Val	Lys	Ser	Tyr	Met
				20					25					30		
	Leu	Glu	Gln	Gly	Ile	Arg	Leu	Ser	Arg	Ala	Lys	Arg	Arg	Phe	Met	Tyr
			35					40					45			
	Lys	Glu	Glu	Ala	Met	Lys	Ala	Leu	Glu	Lys	Met	Ala	Pro	Gln	Thr	Ala
		50					55					60				
55	Gly	Glu	Tyr	Glu	Gly	Thr	Asn	Tyr	Gln	Phe	Lys	Met	Pro	Val	Lys	Val
	65					70					75					80

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Asp Lys His Phe Gly Ser Thr Val Tyr Thr Val Asn Asp Lys Gln Asp
 85 90 95
 5 Lys His Gln Arg Val Val Leu Tyr Ala His Gly Gly Ala Trp Phe Gln
 100 105 110
 Asp Pro Leu Lys Ile His Phe Glu Phe Ile Asp Glu Leu Ala Glu Thr
 115 120 125
 10 Leu Asn Ala Lys Val Ile Met Pro Val Tyr Pro Lys Ile Pro His Gln
 130 135 140
 Asp Tyr Gln Ala Thr Tyr Val Leu Phe Glu Lys Leu Tyr His Asp Leu
 145 150 155 160
 15 Leu Asn Gln Val Ala Asp Ser Lys Gln Ile Val Val Met Gly Asp Ser
 165 170 175
 Ala Gly Gly Gln Ile Ala Leu Ser Phe Ala Gln Leu Leu Lys Glu Lys
 180 185 190
 20 His Ile Val Gln Pro Gly His Ile Val Leu Ile Ser Pro Val Leu Asp
 195 200 205
 Ala Thr Met Gln His Pro Glu Ile Pro Asp Tyr Leu Lys Lys Asp Pro
 210 215 220
 25 Met Val Gly Val Asp Gly Ser Val Phe Leu Ala Glu Gln Trp Ala Gly
 225 230 235 240
 Asp Thr Pro Leu Asp Asn Tyr Lys Val Ser Pro Ile Asn Gly Asp Leu
 245 250 255
 30 Asp Gly Leu Gly Arg Ile Thr Leu Thr Val Gly Thr Lys Glu Val Leu
 260 265 270
 Tyr Pro Asp Ala Leu Asn Leu Ser Gln Leu Leu Ser Ala Lys Gly Ile
 275 280 285
 35 Glu His Asp Phe Ile Pro Gly Tyr Tyr Gln Phe His Ile Tyr Pro Val
 290 295 300
 Phe Pro Ile Pro Glu Arg Arg Arg Phe Leu Tyr Gln Val Lys Asn Ile
 305 310 315 320
 Ile Asn
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 <211> 143
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 <213> Staphylococcus aureus
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 Leu Phe Ser Ala Asp Phe Thr Tyr Gln Ser Val Glu Gln Thr His Gln
 20 25 30
 50 Ser His Ala Ala Val Asn Tyr Tyr Ser Lys Asn Gln Cys Thr Trp Trp
 35 40 45
 Ala Phe Lys Arg Arg Ala Gln Val Gly Lys Pro Val Ser Asn Arg Trp
 50 55 60
 55 Gly Asn Ala Lys Asn Trp Tyr Tyr Asn Ala Arg Lys Ser Lys Tyr Ala
 65 70 75 80

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Thr Gly Arg Thr Pro Arg Lys Phe Ala Val Met Gln Ser Thr Ala Gly
 85 90 95
 Tyr Tyr Gly His Val Ala Val Val Glu Gln Val Tyr Lys Asn Gly Ser
 5 100 105 110
 Ile Lys Val Ser Glu Tyr Asn Phe Tyr Arg Pro Leu Lys Tyr Asn Thr
 115 120 125
 Arg Val Leu Ser Lys Lys Ala Ala Arg Asn Phe Asn Tyr Ile Tyr
 10 130 135 140
 <210> 125
 <211> 255
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 <213> Staphylococcus aureus
 15
 <400> 125
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 20 20 25 30
 Gly Tyr Asn Ser Asn Asp Ala Gln Ser Tyr Ser Tyr Thr Tyr Thr Ile
 35 40 45
 Asp Ala Gln Gly Asn Tyr His Tyr Thr Trp Thr Gly Asn Trp Asn Pro
 50 55 60
 Ser Gln Leu Thr Gln Asn Asn Thr Tyr Tyr Tyr Asn Asn Tyr Asn Thr
 25 65 70 75 80
 Tyr Ser Tyr Asn Asn Ala Ser Tyr Asn Asn Tyr Tyr Asn His Ser Tyr
 85 90 95
 30 Gln Tyr Asn Asn Tyr Thr Asn Asn Ser Gln Thr Ala Thr Asn Asn Tyr
 100 105 110
 Tyr Thr Gly Gly Ser Gly Ala Ser Tyr Ser Thr Thr Ser Asn Asn Val
 115 120 125
 35 His Val Thr Thr Thr Ala Ala Pro Ser Ser Asn Gly Arg Ser Ile Ser
 130 135 140
 Asn Gly Tyr Ala Ser Gly Ser Asn Leu Tyr Thr Ser Gly Gln Cys Thr
 145 150 155 160
 40 Tyr Tyr Val Phe Asp Arg Val Gly Gly Lys Ile Gly Ser Thr Trp Gly
 165 170 175
 Asn Ala Ser Asn Trp Ala Asn Ala Ala Ala Ser Ser Gly Tyr Thr Val
 180 185 190
 45 Asn Asn Thr Pro Lys Val Gly Ala Ile Met Gln Thr Thr Gln Gly Tyr
 195 200 205
 Tyr Gly His Val Ala Tyr Val Glu Gly Val Asn Ser Asn Gly Ser Val
 210 215 220
 50 Arg Val Ser Glu Met Asn Tyr Gly His Gly Ala Gly Val Val Thr Ser
 225 230 235 240
 Arg Thr Ile Ser Ala Asn Gln Ala Gly Ser Tyr Asn Phe Ile His
 245 250 255
 55 <210> 126
 <211> 131
 <212> PRT
 <213> Staphylococcus aureus

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<400> 126
 Met Lys Lys Leu Ile Ile Ser Ile Met Ala Val Met Leu Phe Leu Thr
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 5 Gly Cys Gly Lys Ser Gln Glu Lys Ala Thr Leu Glu Lys Asp Ile Asp
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 35 40 45
 10 Gln Gln Glu Lys Glu Lys Leu Ala Asp Lys Gln Lys Asp Leu Glu Lys
 50 55 60
 Glu Val Lys Asp Leu Lys Pro Ser Lys Glu Asp Asn Lys Asp Asp Lys
 65 70 75 80
 15 Lys Asp Glu Asp Lys Asn Lys Asp Lys Asp Lys Asp Lys Glu Ala Ser
 85 90 95
 Gln Asp Lys Gln Ser Lys Asp Gln Thr Lys Ser Ser Asp Lys Asp Asn
 100 105 110
 20 His Lys Lys Pro Thr Ser Ala Asp Lys Asp Gln Lys Ala Asn Asp Lys
 115 120 125
 His Gln Ser
 130
 25 <210> 127
 <211> 192
 <212> PRT
 <213> Staphylococcus aureus
 <400> 127
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 30 Ser Leu Leu Phe Thr Ile Ile Tyr Ile Asp Asp Ile Gln Lys Trp Phe
 20 25 30
 Asn Gln Tyr Thr Asp Lys Leu Thr Gln Asn His Lys Gly Gln Gly His
 35 40 45
 Ser Lys Trp Glu Asp Phe Phe Arg Gly Ser Arg Ile Thr Glu Thr Phe
 50 55 60
 40 Gly Lys Tyr Gln His Ser Pro Phe Asp Gly Lys His Tyr Gly Ile Asp
 65 70 75 80
 Phe Ala Leu Pro Lys Gly Thr Pro Ile Lys Ala Pro Thr Asn Gly Lys
 85 90 95
 45 Val Thr Arg Ile Phe Asn Asn Glu Leu Gly Gly Lys Val Leu Gln Ile
 100 105 110
 Ala Glu Asp Asn Gly Glu Tyr His Gln Trp Tyr Leu His Leu Asp Lys
 115 120 125
 Tyr Asn Val Lys Val Gly Asp Arg Val Lys Ala Gly Asp Ile Ile Ala
 130 135 140
 50 Tyr Ser Gly Asn Thr Gly Ile Gln Thr Thr Gly Ala His Leu His Phe
 145 150 155 160
 Gln Arg Met Lys Gly Gly Val Gly Asn Ala Tyr Ala Glu Asp Pro Lys
 165 170 175
 55 Pro Phe Ile Asp Gln Leu Pro Asp Gly Glu Arg Ser Leu Tyr Asp Leu
 180 185 190

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<210> 128
 <211> 505
 <212> PRT
 <213> Staphylococcus aureus

5
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 10 20 25 30
 Ile Thr Gly Ala Leu Val Gly Ser Ala Leu Gly Leu Tyr Phe Lys Asn
 35 40 45
 Lys Val Tyr Gln Lys Ala Asp Asp Leu Lys Val Lys Glu Gln Glu Leu
 50 55 60
 Ser Gln Lys Phe Glu Glu Arg Lys Thr Gln Leu Glu Glu Thr Val Ala
 65 70 75 80
 Tyr Thr Lys Glu Arg Val Glu Gly Phe Leu Asn Lys Ser Lys Asn Glu
 85 90 95
 Gln Ala Ala Leu Lys Ala Gln Gln Ala Ala Ile Lys Glu Glu Ala Ser
 100 105 110
 Ala Asn Asn Leu Ser Asp Thr Ser Gln Glu Ala Gln Glu Ile Gln Glu
 115 120 125
 Ala Lys Arg Glu Ala Gln Ala Glu Ala Asp Lys Ser Val Ala Val Ser
 130 135 140
 Asn Lys Glu Ser Lys Ala Val Ala Leu Lys Ala Gln Gln Ala Ala Ile
 145 150 155 160
 Lys Glu Glu Ala Ser Ala Asn Asn Leu Ser Asp Thr Ser Gln Glu Ala
 165 170 175
 Gln Glu Ile Gln Glu Ala Lys Lys Glu Ala Gln Ala Glu Thr Asp Lys
 180 185 190
 Ser Ala Ala Val Ser Asn Glu Glu Pro Lys Ala Val Ala Leu Lys Ala
 195 200 205
 Gln Gln Ala Ala Ile Lys Glu Glu Ala Ser Ala Asn Asn Leu Ser Asp
 210 215 220
 Thr Ser Gln Glu Ala Gln Glu Val Gln Glu Ala Lys Lys Glu Ala Gln
 225 230 235 240
 Ala Glu Thr Asp Lys Ser Ala Ala Val Ser Asn Glu Glu Pro Lys Ala
 245 250 255
 Val Ala Leu Lys Ala Gln Gln Ala Ala Ile Lys Glu Glu Ala Ser Ala
 260 265 270
 Asn Asn Leu Ser Asp Ile Ser Gln Glu Ala Gln Glu Val Gln Glu Ala
 275 280 285
 Lys Lys Glu Ala Gln Ala Glu Lys Asp Ser Asp Thr Leu Thr Lys Asp
 290 295 300
 Ala Ser Ala Ala Lys Val Glu Val Ser Lys Pro Glu Ser Gln Ala Glu
 305 310 315 320
 Arg Leu Ala Asn Ala Ala Lys Gln Lys Gln Ala Lys Leu Thr Pro Gly
 325 330 335

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 Asp Val Ser Glu Thr Glu Thr Val Asn Ile Asp Asn Lys Asp Thr Val
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 Pro Ala Glu Lys Asp Val Val Glu Phe Ala Lys Lys His Lys Asp Lys
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 Ile Ala Lys Arg Gly Glu Gln Phe Phe Met Asp Asn Phe Gly Leu Lys
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 Val Lys Ala Thr Asn Val Val Gly Ser Gly Lys Gly Val Glu Val Phe
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 Val His Cys Asp Asp His Asp Ile Val Phe Asn Ala Ser Ile Pro Phe
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 Asp Lys Ser Ile Ile Glu Ser Asp Ser Ser Leu Arg Ser Glu Asp Lys
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 Asp Asn Glu Glu Lys Tyr Gln Tyr Thr Gly Phe Thr Lys Glu Ala Ile

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	Pro	Ile 50	Leu	Glu	Arg	Lys	Asn 55	Val	Thr	Gly	Phe	Lys 60	Tyr	Thr	Asp	Glu
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	Thr	Leu	Leu	Gly	Ser 85	Asp	Lys	Asp	Lys	Phe 90	Lys	Asp	Gly	Glu	Asn 95	Ser
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	Asn	Tyr	Ser 115	Ile	Gly	Gly	Val	Thr 120	Lys	Ser	Asn	Ser	Val 125	Gln	Tyr	Ile
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	Val 145	Leu	Lys	Asp	Phe	Tyr 150	Tyr	Ile	Ser	Lys	Glu 155	Asp	Ile	Ser	Leu	Lys 160
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	Tyr	Ser	Asn	Gly 180	Leu	Lys	Gln	Gly	Gln 185	Ile	Thr	Ile	Thr	Met 190	Asn	Asp

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 35 40 45
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 85 90 95
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 Glu Gly Ala Lys Tyr Ser Ile Gly Gly Ile Thr Ser Ala Asn Asp Lys
 115 120 125
 35 Ala Val Asp Leu Ile Ala Glu Ala Arg Val Ile Lys Glu Asp His Thr
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 Gly Glu Tyr Asp Tyr Asp Phe Phe Pro Phe Lys Ile Asp Lys Glu Ala
 145 150 155 160
 40 Met Ser Leu Lys Glu Ile Asp Phe Lys Leu Arg Lys Tyr Leu Ile Asp
 165 170 175
 Asn Tyr Gly Leu Tyr Gly Glu Met Ser Thr Gly Lys Ile Thr Val Lys
 180 185 190
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 10 65 70 75 80
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 85 90 95
 Gln Pro Lys Gln Glu Glu Gln Lys Thr Leu Asn Ile Ser Ala Thr Pro
 100 105 110
 Ala Pro Lys Gln Glu Gln Ser Gln Thr Thr Thr Glu Ser Thr Thr Pro
 115 120 125
 Lys Thr Lys Val Thr Thr Pro Pro Ser Thr Asn Thr Pro Gln Pro Met
 130 135 140
 Gln Ser Thr Lys Ser Asp Thr Pro Gln Ser Pro Thr Ile Lys Gln Ala
 145 150 155 160
 Gln Thr Asp Met Thr Pro Lys Tyr Glu Asp Leu Arg Ala Tyr Tyr Thr
 165 170 175
 Lys Pro Ser Phe Glu Phe Glu Lys Gln Phe Gly Phe Met Leu Lys Pro
 180 185 190
 Trp Thr Thr Val Arg Phe Met Asn Val Ile Pro Asn Arg Phe Ile Tyr
 195 200 205
 Lys Ile Ala Leu Val Gly Lys Asp Glu Lys Lys Tyr Lys Asp Gly Pro
 210 215 220
 Tyr Asp Asn Ile Asp Val Phe Ile Val Leu Glu Asp Asn Lys Tyr Gln
 225 230 235 240
 Leu Lys Lys Tyr Ser Val Gly Gly Ile Thr Lys Thr Asn Ser Lys Lys
 245 250 255
 Val Asn His Lys Val Glu Leu Ser Ile Thr Lys Lys Asp Asn Gln Gly
 260 265 270
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 275 280 285
 Ser Leu Lys Glu Leu Asp Phe Lys Leu Arg Lys Gln Leu Ile Glu Lys
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 His Asn Leu Tyr Gly Asn Met Gly Ser Gly Thr Ile Val Ile Lys Met
 305 310 315 320
 Lys Asn Gly Gly Lys Tyr Thr Phe Glu Leu His Lys Lys Leu Gln Glu
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 10 Lys Ile Glu Ala Pro Gln Ser Lys Pro Asn Ala Thr Thr Pro Pro Ser
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 15 Ser Thr Lys Val Thr Thr Pro Pro Ser Thr Asn Thr Pro Gln Pro Met
 85 90 95
 Gln Ser Thr Lys Ser Asp Thr Pro Gln Ser Pro Thr Thr Lys Gln Val
 100 105
 20 Pro Thr Glu Ile Asn Pro Lys Phe Lys Asp Leu Arg Ala Tyr Tyr Thr
 115 120 125
 Lys Pro Ser Leu Glu Phe Lys Asn Glu Ile Gly Ile Ile Leu Lys Lys
 130 135 140
 25 Trp Thr Thr Ile Arg Phe Met Asn Val Val Pro Asp Tyr Phe Ile Tyr
 145 150 155 160
 Lys Ile Ala Leu Val Gly Lys Asp Asp Lys Lys Tyr Gly Glu Gly Val
 165 170 175
 30 His Arg Asn Val Asp Val Phe Val Val Leu Glu Glu Asn Asn Tyr Asn
 180 185 190
 Leu Glu Lys Tyr Ser Val Gly Gly Ile Thr Lys Ser Asn Ser Lys Lys
 195 200 205
 35 Val Asp His Lys Ala Gly Val Arg Ile Thr Lys Glu Asp Asn Lys Gly
 210 215 220
 Thr Ile Ser His Asp Val Ser Glu Phe Lys Ile Thr Lys Glu Gln Ile
 225 230 235 240
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 245 250 255
 Asn Asn Leu Tyr Gly Asn Val Gly Ser Gly Lys Ile Val Ile Lys Met
 260 265 270
 45 Lys Asn Gly Gly Lys Tyr Thr Phe Glu Leu His Lys Lys Leu Gln Glu
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5 His Lys Ala Lys Tyr Glu Asn Val Thr Lys Asp Ile Phe Asp Leu Arg
35 40 45

Asp Tyr Tyr Ser Gly Ala Ser Lys Glu Leu Lys Asn Val Thr Gly Tyr
50 55 60

10 Arg Tyr Ser Lys Gly Gly Lys His Tyr Leu Ile Phe Asp Lys Asn Arg
65 70 75 80

Lys Phe Thr Arg Val Gln Ile Phe Gly Lys Asp Ile Glu Arg Phe Lys
85 90 95

15 Ala Arg Lys Asn Pro Gly Leu Asp Ile Phe Val Val Lys Glu Ala Glu
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Asn Arg Asn Gly Thr Val Phe Ser Tyr Gly Gly Val Thr Lys Lys Asn
115 120 125

20 Gln Asp Ala Tyr Tyr Asp Tyr Ile Asn Ala Pro Arg Phe Gln Ile Lys
130 135 140

Arg Asp Glu Gly Asp Gly Ile Ala Thr Tyr Gly Arg Val His Tyr Ile
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25 Tyr Lys Glu Glu Ile Ser Leu Lys Glu Leu Asp Phe Lys Leu Arg Gln
165 170 175

Tyr Leu Ile Gln Asn Phe Asp Leu Tyr Lys Lys Phe Pro Lys Asp Ser
180 185 190

30 Lys Ile Lys Val Ile Met Lys Asp Gly Gly Tyr Tyr Thr Phe Glu Leu
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35 40 45

Lys Pro Ser Ile Glu Leu Lys Asn Leu Asp Gly Leu Tyr Arg Gln Lys
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Val Thr Asp Lys Gly Val Tyr Val Trp Lys Asp Arg Lys Asp Tyr Phe
65 70 75 80

Val Gly Leu Leu Gly Lys Asp Ile Glu Lys Tyr Pro Gln Gly Glu His
85 90 95

55 Asp Lys Gln Asp Ala Phe Leu Val Ile Glu Glu Glu Thr Val Asn Gly
100 105 110

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 Ser Lys Glu Val Asp Val Lys Val Thr Arg Lys Ile Asp Glu Ser Ser
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 Leu Lys Glu Leu Asp Phe Lys Leu Arg Lys Lys Leu Met Glu Glu Glu
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 Lys Leu Tyr Gly Ala Val Asn Asn Arg Lys Gly Lys Ile Val Val Lys
 180 185 190
 Met Glu Asp Asp Lys Phe Tyr Thr Phe Glu Leu Thr Lys Lys Leu Gln
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 Lys Gln Glu Arg Val Gln His Leu Tyr Asp Ile Lys Asp Leu His Arg
 35 40 45
 Tyr Tyr Ser Ser Glu Ser Phe Glu Phe Ser Asn Ile Ser Gly Lys Val
 50 55 60
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 Glu Asn Tyr Asn Gly Ser Ile Asn Val Val Arg Phe Asn Gln Glu Asn Gln
 65 70 75 80
 Asn His Gln Leu Phe Leu Leu Gly Lys Asp Lys Glu Lys Tyr Lys Glu
 85 90 95
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 Gly Ile Glu Gly Lys Asp Val Phe Val Val Lys Glu Leu Ile Asp Pro
 100 105 110
 Asn Gly Arg Leu Ser Thr Val Gly Gly Val Thr Lys Lys Asn Asn Lys
 115 120 125
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 Ser Ser Glu Thr Asn Thr His Leu Phe Val Asn Lys Val Tyr Gly Gly
 130 135 140
 Asn Leu Asp Ala Ser Ile Asp Ser Phe Ser Ile Asn Lys Glu Glu Val
 145 150 155 160
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 165 170 175
 Tyr Gly Leu Tyr Lys Gly Thr Thr Lys Tyr Gly Lys Ile Thr Ile Asn
 180 185 190
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 35 40 45
 Tyr Tyr Ser Gly Pro Ser Tyr Glu Leu Thr Asn Val Ser Gly Gln Ser
 50 60
 Gln Gly Tyr Tyr Asp Ser Asn Val Leu Leu Phe Asn Gln Gln Asn Gln
 65 70 75 80
 Lys Phe Gln Val Phe Leu Leu Gly Lys Asp Glu Asn Lys Tyr Lys Glu
 85 90 95
 Lys Thr His Gly Leu Asp Val Phe Ala Val Pro Glu Leu Val Asp Leu
 100 105 110
 Asp Gly Arg Ile Phe Ser Val Ser Gly Val Thr Lys Lys Asn Val Lys
 115 120 125
 Ser Ile Phe Glu Ser Leu Arg Thr Pro Asn Leu Leu Val Lys Lys Ile
 130 135 140
 Asp Asp Lys Asp Gly Phe Ser Ile Asp Glu Phe Phe Phe Ile Gln Lys
 145 150 155 160
 Glu Glu Val Ser Leu Lys Glu Leu Asp Phe Lys Ile Arg Lys Leu Leu
 165 170 175
 Ile Lys Lys Tyr Lys Leu Tyr Glu Gly Ser Ala Asp Lys Gly Arg Ile
 180 185 190
 Val Ile Asn Met Lys Asp Glu Asn Lys Tyr Glu Ile Asp Leu Ser Asp
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 35 40 45

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Tyr Tyr Ser Glu Glu Ser Phe Glu Pro Thr Asn Ile Ser Val Lys Ser
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 Glu Asp Tyr Tyr Gly Ser Asn Val Leu Asn Phe Lys Gln Arg Asn Lys
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 Ala Phe Lys Val Phe Leu Leu Gly Asp Asp Lys Asn Lys Tyr Lys Glu
 85 90 95
 Lys Thr His Gly Leu Asp Val Phe Ala Val Pro Glu Leu Ile Asp Ile
 10 100 105 110
 Lys Gly Gly Ile Tyr Ser Val Gly Gly Ile Thr Lys Lys Asn Val Arg
 115 120 125
 Ser Val Phe Gly Phe Val Ser Asn Pro Ser Leu Gln Val Lys Lys Val
 15 130 135 140
 Asp Ala Lys Asn Gly Phe Ser Ile Asn Glu Leu Phe Phe Ile Gln Lys
 145 150 155 160
 Glu Glu Val Ser Leu Lys Glu Leu Asp Phe Lys Ile Arg Lys Leu Leu
 20 165 170 175
 Ile Glu Lys Tyr Arg Leu Tyr Lys Gly Thr Ser Asp Lys Gly Arg Ile
 180 185 190
 Val Ile Asn Met Lys Asp Glu Lys Lys His Glu Ile Asp Leu Ser Glu
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 Tyr Thr Gly Lys Thr Met Glu Met Lys Asn Ile Ser Ala Leu Lys His
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 Gly Lys Asn Asn Leu Arg Phe Lys Phe Arg Gly Ile Lys Ile Gln Val
 65 70 75 80
 Leu Leu Pro Gly Asn Asp Lys Ser Lys Phe Gln Gln Arg Ser Tyr Glu
 85 90 95
 Gly Leu Asp Val Phe Phe Val Gln Glu Lys Arg Asp Lys His Asp Ile
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 Phe Tyr Thr Val Gly Gly Val Ile Gln Asn Asn Lys Thr Ser Gly Val
 115 120 125
 55 Val Ser Ala Pro Ile Leu Asn Ile Ser Lys Glu Lys Gly Glu Asp Ala
 130 135 140

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Phe Val Lys Gly Tyr Pro Tyr Tyr Ile Lys Lys Glu Lys Ile Thr Leu
 145 150 155 160
 5 Lys Glu Leu Asp Tyr Lys Leu Arg Lys His Leu Ile Glu Lys Tyr Gly
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 Leu Tyr Lys Thr Ile Ser Lys Asp Gly Arg Val Lys Ile Ser Leu Lys
 180 185 190
 10 Asp Gly Ser Phe Tyr Asn Leu Asp Leu Arg Ser Lys Leu Lys Phe Lys
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 50 55 60
 30 Lys Val Thr Phe Thr Pro Asn Tyr Gln Leu Ile Asp Val Thr Leu Thr
 65 70 75 80
 Gly Asn Glu Lys Gln Asn Phe Gly Glu Asp Ile Ser Asn Val Asp Ile
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 35 Phe Val Val Arg Glu Asn Ser Asp Arg Ser Gly Asn Thr Ala Ser Ile
 100 105 110
 Gly Gly Ile Thr Lys Thr Asn Gly Ser Asn Tyr Ile Asp Lys Val Lys
 115 120 125
 40 Asp Val Asn Leu Ile Ile Thr Lys Asn Ile Asp Ser Val Thr Ser Thr
 130 135 140
 Ser Thr Ser Ser Thr Tyr Thr Ile Asn Lys Glu Glu Ile Ser Leu Lys
 145 150 155 160
 45 Glu Leu Asp Phe Lys Leu Arg Lys His Leu Ile Asp Lys His Asn Leu
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 Tyr Lys Thr Glu Pro Lys Asp Ser Lys Ile Arg Ile Thr Met Lys Asp
 180 185 190
 50 Gly Gly Phe Tyr Thr Phe Glu Leu Asn Lys Lys Leu Gln Thr His Arg
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 35 40 45
 Ala Glu Glu Leu Arg Thr Leu Leu Asn Lys Ser Asn Val Tyr Ala Leu
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 65 70 75 80
 Glu Tyr Arg Ala Lys Ala Ala Leu Lys Lys Asn Asp Phe Val Ser Met
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 35 40 45
 Ser Arg Thr Val Glu Glu Tyr Lys Lys Ser Leu Asp Asp Leu Ile Trp
 50 55 60
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 65 70 75 80
 Glu Ala Met Lys Lys Tyr Gln Gln Arg Phe Met Ala Glu Asp Glu Ala
 85 90 95
 45 Leu Lys Lys Phe Phe Ser Glu Glu Lys Lys Ile Lys Asn Gly Asn Thr
 100 105 110
 Asp Asn Leu Asp Tyr Leu Gly Leu Ser His Glu Arg Tyr Glu Ser Val
 115 120 125
 50 Phe Asn Thr Leu Lys Lys Gln Ser Glu Glu Phe Leu Lys Glu Ile Glu
 130 135 140
 55 Asp Ile Lys Lys Asp Asn Pro Glu Leu Lys Asp Phe Asn Glu Glu Glu
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 Gln Leu Lys Cys Asp Leu Glu Leu Asn Lys Leu Glu Asn Gln Ile Leu
 165 170 175
 Met Leu Gly Lys Thr Phe Tyr Gln Asn Tyr Arg Asp Asp Val Glu Ser
 180 185 190

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Leu Tyr Ser Lys Leu Asp Leu Ile Met Gly Tyr Lys Asp Glu Glu Arg
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 Asp Leu Glu Thr Ile Ile Asp Glu Phe Phe Ser Asp Ile Asp Lys Thr
 225 230 235 240
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 260 265 270
 15 Ser Asp Glu Ser Lys Arg Ser Lys Arg Ser Lys Arg Ser Leu Asn Thr
 275 280 285
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 20 Glu Tyr Asp Lys Arg Ala Glu Glu Arg Lys Ala Arg Phe Leu Asp Asn
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 325 330 335
 25 His Lys Gln Arg Ile Asp Asn Glu Asn Asp Lys Lys Leu Val Val Ser
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 30 Thr Gln Val Pro Met Pro Thr Val Glu Arg Gln Thr Gln Gln Gln Ile
 370 375 380
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 35 Asp Phe Thr Thr Thr His Gln Ser Pro Thr Thr Ser Asn His Thr His
 405 410 415
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 420 425 430
 40 Ser Gly Ser Leu Val Gly Ile Ser Gln Ile Asp Ser Ser His Leu Thr
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 Glu Arg Glu Lys Arg Val Ile Lys Arg Glu His Val Arg Glu Ala Gln
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 45 Lys Leu Val Asp Asn Tyr Lys Asp Thr His Ser Tyr Lys Asp Arg Ile
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 50 Asn Ala Gln Gln Lys Val Asn Thr Leu Ser Glu Gly His Gln Lys Arg
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 5 Gln Val Asp Lys Ile Lys Asp Asp Glu Glu Pro Ile Lys Thr Val Gly
 35 40 45
 Lys Lys Ile Ala Glu Leu Asp Glu Lys Lys Lys Lys Leu Thr Glu Asp
 50 55 60
 10 Val Asn Ser Lys Asp Thr Ala Val Arg Gly Lys Ala Val Lys Asp Leu
 65 70 75 80
 Ile Lys Asn Ala Asp Asp Arg Leu Lys Glu Phe Glu Lys Glu Glu Asp
 85 90 95
 15 Ala Ile Lys Lys Ser Glu Gln Asp Phe Lys Lys Ala Lys Ser His Val
 100 105 110
 Asp Asn Ile Asp Asn Asp Val Lys Arg Lys Glu Val Lys Gln Leu Asp
 115 120 125
 20 Asp Val Leu Lys Glu Lys Tyr Lys Leu His Ser Asp Tyr Ala Lys Ala
 130 135 140
 Tyr Lys Lys Ala Val Asn Ser Glu Lys Thr Leu Phe Lys Tyr Leu Asn
 145 150 155 160
 25 Gln Asn Asp Ala Thr Gln Gln Gly Val Asn Glu Lys Ser Lys Ala Ile
 165 170 175
 Glu Gln Asn Tyr Lys Lys Leu Lys Glu Val Ser Asp Lys Tyr Thr Lys
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 30 Val Leu Asn Lys Val Gly Lys Glu Lys Gln Asp Val Asp Gln Phe Lys
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 Ser Leu Thr Val Ser Thr Phe Ala Gly Glu Ser His Ala Gln Thr Lys
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 40 Asn Val Glu Ala Ala Lys Lys Tyr Asp Gln Tyr Gln Thr Asn Phe Lys
 35 40 45
 Lys Gln Val Asn Lys Lys Val Val Asp Ala Gln Lys Ala Val Asn Phe
 50 55 60
 45 Phe Lys Arg Thr Arg Thr Val Ala Thr His Arg Lys Ala Gln Arg Ala
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 5 Thr Ser Leu Pro Thr Ser Asn Glu Tyr Gln Asn Glu Lys Leu Ala Asn
 35 40 45
 Glu Leu Lys Ser Leu Leu Asp Glu Leu Asn Val Asn Glu Leu Ala Thr
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 10 Gly Ser Leu Asn Thr Tyr Tyr Lys Arg Thr Ile Lys Ile Ser Gly Gln
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 Lys Ala Met Tyr Ala Leu Lys Ser Lys Asp Phe Lys Lys Met Ser Glu
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Glu Lys Asp Leu Arg Lys Lys Ser Glu Leu Gln Gly Ala Ala Leu Gly
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 5 Asn Leu Lys Gln Ile Tyr Tyr Tyr Asn Glu Lys Ala Lys Thr Glu Asn
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 Lys Glu Ser His Asp Gln Phe Leu Gln His Thr Ile Leu Phe Lys Gly
 65 70 75 80
 10 Phe Phe Thr Asn His Ser Trp Tyr Asn Asp Leu Leu Val Asp Phe Asp
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 Ser Lys Asp Ile Val Asp Lys Tyr Lys Gly Lys Lys Val Asp Leu Tyr
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 15 Gly Ala Tyr Tyr Gly Tyr Gln Cys Ala Gly Gly Thr Pro Asn Lys Thr
 115 120 125
 Ala Cys Met Tyr Gly Gly Val Thr Leu His Asp Asn Asn Arg Leu Thr
 130 135 140
 20 Glu Glu Lys Lys Val Pro Ile Asn Leu Trp Leu Asp Gly Lys Gln Asn
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 Thr Val Pro Leu Glu Thr Val Lys Thr Asn Lys Lys Asn Val Thr Val
 165 170 175
 25 Gln Glu Leu Asp Leu Gln Ala Arg Arg Tyr Leu Gln Glu Lys Tyr Asn
 180 185 190
 Leu Tyr Asn Ser Asp Val Phe Asp Gly Lys Val Gln Arg Gly Leu Ile
 195 200 205
 30 Val Phe His Thr Ser Thr Glu Pro Ser Val Asn Tyr Asp Leu Phe Gly
 210 215 220
 Ala Gln Gly Gln Asn Ser Asn Thr Leu Leu Arg Ile Tyr Arg Asp Asn
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 35 40 45
 50 Asp Lys Glu Lys Glu Val Ala Thr Gln Gln Gln Pro Asp Asn Gln Thr
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 Val Glu Gln Pro Gln Ser Gln Glu Gln Ser Val Gln Gln Pro Gln Gln
 65 70 75 80
 55 Gln Ile Pro Gln Asn Ser Val Pro Gln Gln Asn Val Gln Val Gln Gln
 85 90 95

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 Ser Thr Glu Gly Met Ser Glu Gln Ala Gln Lys Gln Ile Glu Glu Leu
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 35 40 45
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 25 Asp Asn Val Gln Ser Lys Glu Val Lys Ile Glu Glu Val Thr Asn Lys
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 Asp Thr Ala Pro Gln Gly Val Glu Ala Lys Ser Glu Val Thr Ser Asn
 85 90 95
 30 Lys Asp Thr Ile Glu His Glu Pro Ser Val Lys Ala Glu Asp Ile Ser
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 35 Pro Lys Ser Ser Val Thr His Asn Ala Glu Thr Pro Lys Val Arg Lys
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 Ala Arg Ser Val Asp Glu Gly Ser Phe Asp Ile Thr Arg Asp Ser Lys
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 40 Asn Val Val Glu Ser Thr Pro Ile Thr Ile Gln Gly Lys Glu His Phe
 165 170 175
 Glu Gly Tyr Gly Ser Val Asp Ile Gln Lys Lys Pro Thr Asp Leu Gly
 180 185 190
 45 Val Ser Glu Val Thr Arg Phe Asn Val Gly Asn Glu Ser Asn Gly Leu
 195 200 205
 Ile Gly Ala Leu Gln Leu Lys Asn Lys Ile Asp Phe Ser Lys Asp Phe
 210 215 220
 50 Asn Phe Lys Val Arg Val Ala Asn Asn His Gln Ser Asn Thr Thr Gly
 225 230 235 240
 Ala Asp Gly Trp Gly Phe Leu Phe Ser Lys Gly Asn Ala Glu Glu Tyr
 245 250 255
 55 Leu Thr Asn Gly Gly Ile Leu Gly Asp Lys Gly Leu Val Asn Ser Gly
 260 265 270
 Gly Phe Lys Ile Asp Thr Gly Tyr Ile Tyr Thr Ser Ser Met Asp Lys

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15	Trp	Glu	Thr	Ser	Ile	Thr	Asp	Leu	Gly	Leu	Ser	Lys	Asn	Gln	Ala	Tyr
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	625					630					635					640
	Asp	Pro	Lys	Leu	Pro	Thr	Gly	Glu	Lys	Glu	Glu	Val	Pro	Gly	Lys	Pro
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Gly Ile Lys Asn Pro Glu Thr Gly Asp Val Val Arg Pro Pro Val Asp
 660 665 670
 Ser Val Thr Lys Tyr Gly Pro Val Lys Gly Asp Ser Ile Val Glu Lys
 5 675 680 685
 Glu Glu Ile Pro Phe Glu Lys Glu Arg Lys Phe Asn Pro Asp Leu Ala
 690 695 700
 Pro Gly Thr Glu Lys Val Thr Arg Glu Gly Gln Lys Gly Glu Lys Thr
 10 705 710 715 720
 Ile Thr Thr Pro Thr Leu Lys Asn Pro Leu Thr Gly Val Ile Ile Ser
 725 730 735
 Lys Gly Glu Pro Lys Glu Glu Ile Thr Lys Asp Pro Ile Asn Glu Leu
 15 740 745 750
 Thr Glu Tyr Gly Pro Glu Thr Ile Thr Pro Gly His Arg Asp Glu Phe
 755 760 765
 Asp Pro Lys Leu Pro Thr Gly Glu Lys Glu Glu Val Pro Gly Lys Pro
 20 770 775 780
 Gly Ile Lys Asn Pro Glu Thr Gly Asp Val Val Arg Pro Pro Val Asp
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 Ser Val Thr Lys Tyr Gly Pro Val Lys Gly Asp Ser Ile Val Glu Lys
 25 805 810 815
 Glu Glu Ile Pro Phe Lys Lys Glu Arg Lys Phe Asn Pro Asp Leu Ala
 820 825 830
 Pro Gly Thr Glu Lys Val Thr Arg Glu Gly Gln Lys Gly Glu Lys Thr
 30 835 840 845
 Ile Thr Thr Pro Thr Leu Lys Asn Pro Leu Thr Gly Glu Ile Ile Ser
 850 855 860
 Lys Gly Glu Ser Lys Glu Glu Ile Thr Lys Asp Pro Ile Asn Glu Leu
 35 865 870 875 880
 Thr Glu Tyr Gly Pro Glu Thr Ile Thr Pro Gly His Arg Asp Glu Phe
 885 890 895
 Asp Pro Lys Leu Pro Thr Gly Glu Lys Glu Glu Val Pro Gly Lys Pro
 40 900 905 910
 Gly Ile Lys Asn Pro Glu Thr Gly Asp Val Val Arg Pro Pro Val Asp
 915 920 925
 Ser Val Thr Lys Tyr Gly Pro Val Lys Gly Asp Ser Ile Val Glu Lys
 45 930 935 940
 Glu Glu Ile Pro Phe Lys Lys Glu Arg Lys Phe Asn Pro Asp Leu Ala
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 Pro Gly Thr Glu Lys Val Thr Arg Glu Gly Gln Lys Gly Glu Lys Thr
 965 970 975
 50 Ile Thr Thr Pro Thr Leu Lys Asn Pro Leu Thr Gly Glu Ile Ile Ser
 980 985 990
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 55 Thr Glu Tyr Gly Pro Glu Thr Ile Thr Pro Gly His Arg Asp Glu Phe
 1010 1015 1020

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Asp Pro Lys Leu Pro Thr Gly Glu Lys Glu Glu Val Pro Gly Lys Pro
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 5 1045 1050 1055
 Ser Val Thr Lys Tyr Gly Pro Val Lys Gly Asp Ser Ile Val Glu Lys
 1060 1065 1070
 Glu Glu Ile Pro Phe Glu Lys Glu Arg Lys Phe Asn Pro Asp Leu Ala
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 Pro Gly Thr Glu Lys Val Thr Arg Glu Gly Gln Lys Gly Glu Lys Thr
 1090 1095 1100
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 Thr Glu Tyr Gly Pro Glu Thr Ile Thr Pro Gly His Arg Asp Glu Phe
 20 1140 1145 1150
 Asp Pro Lys Leu Pro Thr Gly Glu Lys Glu Glu Val Pro Gly Lys Pro
 1155 1160 1165
 Gly Ile Lys Asn Pro Glu Thr Gly Asp Val Val Arg Pro Pro Val Asp
 25 1170 1175 1180
 Ser Val Thr Lys Tyr Gly Pro Val Lys Gly Asp Ser Ile Val Glu Lys
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 Lys Gly Glu Ser Lys Glu Glu Ile Thr Lys Asp Pro Val Asn Glu Leu
 35 1250 1255 1260
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 Asp Pro Asn Leu Pro Thr Asp Gln Thr Glu Lys Val Pro Gly Lys Pro
 40 1285 1290 1295
 Gly Ile Lys Asn Pro Asp Thr Gly Lys Val Ile Glu Glu Pro Val Asp
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 1330 1335 1340
 Pro Gly Glu Glu Arg Val Lys Gln Glu Gly Gln Pro Gly Ser Lys Thr
 50 1345 1350 1355 1360
 Ile Thr Thr Pro Ile Thr Val Asn Pro Leu Thr Gly Glu Lys Val Gly
 1365 1370 1375
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 55 1380 1385 1390
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10	Glu	Ser	Val	Ala	Asn	Gln	Glu	Lys	Lys	Arg	Ala	Glu	Leu	Pro	Lys	Thr			
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	Gly	Leu	Glu	Ser	Thr	Gln	Lys	Gly	Leu	Ile	Phe	Ser	Ser	Ile	Ile	Gly			
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	Gly	Met	Leu	Lys	Lys	Val	Phe	Tyr	Ser	Phe	Ile	Asp	Asp	Lys	Asn	His			
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	Tyr	Arg	Val	Tyr	Ser	Glu	Glu	Gly	Ala	Asn	Lys	Ser	Gly	Leu	Ala	Trp			
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35	Pro	Ser	Ala	Phe	Lys	Val	Gln	Leu	Gln	Leu	Pro	Asp	Asn	Glu	Val	Ala			
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	Gln	Ile	Ser	Asp	Tyr	Tyr	Pro	Arg	Asn	Ser	Ile	Asp	Thr	Lys	Glu	Tyr			
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40	Met	Ser	Thr	Leu	Thr	Tyr	Gly	Phe	Asn	Gly	Asn	Val	Thr	Gly	Asp	Asp			
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45	Thr	Leu	Lys	Tyr	Val	Gln	Pro	Asp	Phe	Lys	Thr	Ile	Leu	Glu	Ser	Pro			
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	Thr	Asp	Lys	Lys	Val	Gly	Trp	Lys	Val	Ile	Phe	Asn	Asn	Met	Val	Asn			
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50	Gln	Asn	Trp	Gly	Pro	Tyr	Asp	Arg	Asp	Ser	Trp	Asn	Pro	Val	Tyr	Gly			
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	Asn	Gln	Leu	Phe	Met	Lys	Thr	Arg	Asn	Gly	Ser	Met	Lys	Ala	Ala	Asp			
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Gln Gln Thr Asn Ile Asp Val Ile Tyr Glu Arg Val Arg Asp Asp Tyr
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 5 Gln Leu His Trp Thr Ser Thr Asn Trp Lys Gly Thr Asn Thr Lys Asp
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 35 40 45
 25 Ser Arg Phe Glu Glu Gln Phe Gln Gln Leu Ser Pro Lys Val Glu Lys
 50 55 60
 Phe Ala Gln Leu Leu Glu Glu Ile Lys Gln Gln Leu Asn Ser Thr Ala
 65 70 75 80
 30 Asp Ala Val Gln Glu Gln Asp Gln Gln Leu Ser Asn Asn Phe Gly Leu
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 Gln Ala Ser Gly Gly Gly Ser Met Gly Gly Tyr Lys Gly Ile Lys Ala
 100 105 110
 35 Asp Gly Gly Lys Val Asp Gln Ala Lys Gln Leu Ala Ala Lys Thr Ala
 115 120 125
 Lys Asp Ile Glu Ala Cys Gln Lys Gln Thr Gln Gln Leu Ala Glu Tyr
 130 135 140
 40 Ile Glu Gly Ser Asp Trp Glu Gly Gln Phe Ala Asn Lys Val Lys Asp
 145 150 155 160
 Val Leu Leu Ile Met Ala Lys Phe Gln Glu Glu Leu Val Gln Pro Met
 165 170 175
 45 Ala Asp His Gln Lys Ala Ile Asp Asn Leu Ser Gln Asn Leu Ala Lys
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 Tyr Asp Thr Leu Ser Ile Lys Gln Gly Leu Asp Arg Val Asn Pro
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Lys Gln Thr Gln Gln Leu Ala Glu Tyr Ile Glu Gly Ser Asp Trp Glu
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 Gly Gln Phe Ala Asn Lys Val Lys Asp Val Leu Leu Ile Met Ala Lys
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 Phe Gln Glu Glu Leu Val Gln Pro Met Ala Asp His Gln Lys Ala Ile
 65 70 75 80
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 10 85 90 95
 Gln Gly Leu Asp Arg Val Asn Pro Ala Ser Gly Gly Gly Ser Met Ala
 100 105
 Met Ile Lys Met Ser Pro Glu Glu Ile Arg Ala Lys Ser Gln Ser Tyr
 15 115 120 125
 Gly Gln Gly Ser Asp Gln Ile Arg Gln Ile Leu Ser Asp Leu Thr Arg
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 Ala Gln Gly Glu Ile Ala Ala Asn Trp Glu Gly Gln Ala Phe Ser Arg
 20 145 150 155 160
 Phe Glu Glu Gln Phe Gln Gln Leu Ser Pro Lys Val Glu Lys Phe Ala
 165 170 175
 Gln Leu Leu Glu Glu Ile Lys Gln Gln Leu Asn Ser Thr Ala Asp Ala
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 35 40 45
 Lys Gly Gln Lys Leu Lys Gly Ile Ser Phe Glu Asn Ser Asn Gly Glu
 40 50 55 60
 Trp Ala Tyr Lys Val Thr Gln Gln Lys Ser Gly Glu Glu Ser Glu Val
 65 70 75 80
 Leu Val Ala Asp Lys Asn Lys Lys Val Ile Asn Lys Lys Thr Glu Lys
 45 85 90 95
 Glu Asp Thr Met Asn Glu Asn Asp Asn Phe Lys Tyr Ser Asp Ala Ile
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 Asp Tyr Lys Lys Ala Ile Lys Glu Gly Gln Lys Glu Phe Asp Gly Asp
 50 115 120 125
 Ile Lys Glu Trp Ser Leu Glu Lys Asp Asp Gly Lys Leu Val Tyr Asn
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165

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 Lys Gly Gln Lys Leu Lys Gly Ile Ser Phe Glu Asn Ser Asn Gly Glu
 50 55 60
 Trp Ala Tyr Lys Val Thr Gln Gln Lys Ser Gly Glu Glu Ser Glu Val
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 35 40 45
 Gln Pro Glu Ala Lys Lys Glu Ser Thr Ser Ser Ser Thr Gln Lys Gln
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 Gln Asn Asn Val Thr Ala Thr Thr Glu Thr Lys Pro Gln Asn Ile Glu
 65 70 75 80
 Lys Glu Asn Val Lys Pro Ser Thr Asp Lys Thr Ala Thr Glu Asp Thr
 85 90 95
 Ser Val Ile Leu Glu Glu Lys Lys Ala Pro Asn Asn Thr Asn Asn Asp
 100 105 110
 Val Thr Thr Lys Pro Ser Thr Ser Glu Pro Ser Thr Ser Glu Ile Gln
 115 120 125
 Thr Lys Pro Thr Thr Pro Gln Glu Ser Thr Asn Ile Glu Asn Ser Gln
 130 135 140
 Pro Gln Pro Thr Pro Ser Lys Val Asp Asn Gln Val Thr Asp Ala Thr
 145 150 155 160
 Asn Pro Lys Glu Pro Val Asn Val Ser Lys Glu Glu Leu Lys Asn Asn
 165 170 175
 Pro Glu Lys Leu Lys Glu Leu Val Arg Asn Asp Ser Asn Thr Asp His
 180 185 190

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Ser Thr Lys Pro Val Ala Thr Ala Pro Thr Ser Val Ala Pro Lys Arg
 195 200 205
 Val Asn Ala Lys Met Arg Phe Ala Val Ala Gln Pro Ala Ala Val Ala
 210 215 220
 Ser Asn Asn Val Asn Asp Leu Ile Lys Val Thr Lys Gln Thr Ile Lys
 225 230 235 240
 Val Gly Asp Gly Lys Asp Asn Val Ala Ala Ala His Asp Gly Lys Asp
 245 250 255
 Ile Glu Tyr Asp Thr Glu Phe Thr Ile Asp Asn Lys Val Lys Lys Gly
 260 265 270
 Asp Thr Met Thr Ile Asn Tyr Asp Lys Asn Val Ile Pro Ser Asp Leu
 275 280 285
 Thr Asp Lys Asn Asp Pro Ile Asp Ile Thr Asp Pro Ser Gly Glu Val
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 Phe Thr Asp Tyr Val Asp Lys Tyr Glu Asp Ile Lys Ser Arg Leu Thr
 325 330 335
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 Asn Leu Thr Phe Ala Thr Ala Gly Lys Glu Thr Ser Gln Asn Val Thr
 355 360 365
 Val Asp Tyr Gln Asp Pro Met Val His Gly Asp Ser Asn Ile Gln Ser
 370 375 380
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 385 390 395 400
 Tyr Val Asn Pro Leu Lys Lys Ser Ala Thr Asn Thr Lys Val Asp Ile
 405 410 415
 Ala Gly Ser Gln Val Asp Asp Tyr Gly Asn Ile Lys Leu Gly Asn Gly
 420 425 430
 Ser Thr Ile Ile Asp Gln Asn Thr Glu Ile Lys Val Tyr Lys Val Asn
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 Ser Asp Gln Gln Leu Pro Gln Ser Asn Arg Ile Tyr Asp Phe Ser Gln
 450 455 460
 Tyr Glu Asp Val Thr Ser Gln Phe Asp Asn Lys Lys Ser Phe Ser Asn
 465 470 475 480
 Asn Val Ala Thr Leu Asp Phe Gly Asp Ile Asn Ser Ala Tyr Ile Ile
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 Lys Val Val Ser Lys Tyr Thr Pro Thr Ser Asp Gly Glu Leu Asp Ile
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 Ala Gln Gly Thr Ser Met Arg Thr Thr Asp Lys Tyr Gly Tyr Tyr Asn
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 Tyr Ala Gly Tyr Ser Asn Phe Ile Val Thr Ser Asn Asp Thr Gly Gly
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 Gly Asp Gly Thr Val Lys Pro Glu Glu Lys Leu Tyr Lys Ile Gly Asp
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Lys Glu Lys Pro
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Asn Glu Thr Thr Ser Asn Gly Asn Lys Leu Ile Glu Lys Glu Ser Val
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Gln Ser Thr Thr Gly Asn Lys Val Glu Val Ser Thr Ala Lys Ser Asp
65 70 75 80

Glu Gln Ala Ser Pro Lys Ser Thr Asn Glu Asp Leu Asn Thr Lys Gln
85 90 95

25

Thr Ile Ser Asn Gln Glu Ala Leu Gln Pro Asp Leu Gln Glu Asn Lys
100 105 110

Ser Val Val Asn Val Gln Pro Thr Asn Glu Glu Asn Lys Lys Val Asp
115 120 125

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Ala Lys Thr Glu Ser Thr Thr Leu Asn Val Lys Ser Asp Ala Ile Lys
130 135 140

Ser Asn Asp Glu Thr Leu Val Asp Asn Asn Ser Asn Ser Asn Asn Glu
145 150 155 160

35

Asn Asn Ala Asp Ile Ile Leu Pro Lys Ser Thr Ala Pro Lys Arg Leu
165 170 175

Asn Thr Arg Met Arg Ile Ala Ala Val Gln Pro Ser Ser Thr Glu Ala
180 185 190

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Lys Asn Val Asn Asp Leu Ile Thr Ser Asn Thr Thr Leu Thr Val Val
195 200 205

Asp Ala Asp Lys Asn Asn Lys Ile Val Pro Ala Gln Asp Tyr Leu Ser
210 215 220

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Leu Lys Ser Gln Ile Thr Val Asp Asp Lys Val Lys Ser Gly Asp Tyr
225 230 235 240

Phe Thr Ile Lys Tyr Ser Asp Thr Val Gln Val Tyr Gly Leu Asn Pro
245 250 255

50

Glu Asp Ile Lys Asn Ile Gly Asp Ile Lys Asp Pro Asn Asn Gly Glu
260 265 270

Thr Ile Ala Thr Ala Lys His Asp Thr Ala Asn Asn Leu Ile Thr Tyr
275 280 285

55

Thr Phe Thr Asp Tyr Val Asp Arg Phe Asn Ser Val Gln Met Gly Ile
290 295 300

Asn Tyr Ser Ile Tyr Met Asp Ala Asp Thr Ile Pro Val Ser Lys Asn

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Ser Leu Thr Thr Lys Val His Ile Val Val Pro Gln Ile Asn Tyr Asn
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 Asn Lys
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Lys Asp Asp Pro Ser Gln Ser Ala Asn Val Leu Gly Glu Ala Gln Lys
35 40 45

Leu Asn Asp Ser Gln Ala Pro Lys Ala Asp Ala Gln Gln Asn Asn Phe
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Asn Lys Asp Gln Gln Ser Ala Phe Tyr Glu Ile Leu Asn Met Pro Asn
65 70 75 80

Leu Asn Glu Ala Gln Arg Asn Gly Phe Ile Gln Ser Leu Lys Asp Asp
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Pro Ser Gln Ser Thr Asn Val Leu Gly Glu Ala Lys Lys Leu Asn Glu
100 105 110

Ser Gln Ala Pro Lys Ala Asp Asn Asn Phe Asn Lys Glu Gln Gln Asn
115 120 125

Ala Phe Tyr Glu Ile Leu Asn Met Pro Asn Leu Asn Glu Glu Gln Arg
130 135 140

Asn Gly Phe Ile Gln Ser Leu Lys Asp Asp Pro Ser Gln Ser Ala Asn
145 150 155 160

Leu Leu Ser Glu Ala Lys Lys Leu Asn Glu Ser Gln Ala Pro Lys Ala
165 170 175

Asp Asn Lys Phe Asn Lys Glu Gln Gln Asn Ala Phe Tyr Glu Ile Leu
180 185 190

His Leu Pro Asn Leu Asn Glu Glu Gln Arg Asn Gly Phe Ile Gln Ser
195 200 205

Leu Lys Asp Asp Pro Ser Gln Ser Ala Asn Leu Leu Ala Glu Ala Lys
210 215 220

Lys Leu Asn Asp Ala Gln Ala Pro Lys Ala Asp Asn Lys Phe Asn Lys
225 230 235 240

Glu Gln Gln Asn Ala Phe Tyr Glu Ile Leu His Leu Pro Asn Leu Thr
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Glu Glu Gln Arg Asn Gly Phe Ile Gln Ser Leu Lys Asp Asp Pro Ser
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			20			
	Asn	Asn	Met	Ile	Glu	Thr
				25		
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		35			40	
	Val	Asp	Ser	Thr	Thr	Lys
		50			55	
	Thr	Thr	Glu	Pro	Ala	Ser
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	Lys	Asn	Gln	Ala	Thr	Asn
				85		
	Glu	Ala	Asn	Ser	Gln	Val
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	Ile	Ala	Thr	Asn	Ser	Glu
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	Ser	Val	Arg	Thr	Arg	Ala
					150	
	Val	Asn	Ala	Ala	Asp	Ala
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	Ile	Lys	Ser	Thr	Asn	Gly
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	Leu	Thr	Lys	Thr	Tyr	Thr
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	Glu	Asn	Ile	Asn	Gly	Gln
			275			
	Lys	Ala	Pro	Lys	Ser	Gly
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	Glu	Met	Phe	Asn	Asn	Lys
						310

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 Val Asp Thr Ala Ser Gly Gln Asn Thr Tyr Lys Gln Thr Val Phe Val
 5 340 345 350
 Asn Pro Lys Gln Arg Val Leu Gly Asn Thr Trp Val Tyr Ile Lys Gly
 355 360 365
 Tyr Gln Asp Lys Ile Glu Glu Ser Ser Gly Lys Val Ser Ala Thr Asp
 10 370 375 380
 Thr Lys Leu Arg Ile Phe Glu Val Asn Asp Thr Ser Lys Leu Ser Asp
 385 390 395 400
 Ser Tyr Tyr Ala Asp Pro Asn Asp Ser Asn Leu Lys Glu Val Thr Asp
 15 405 410 415
 Gln Phe Lys Asn Arg Ile Tyr Tyr Glu His Pro Asn Val Ala Ser Ile
 420 425 430
 Lys Phe Gly Asp Ile Thr Lys Thr Tyr Val Val Leu Val Glu Gly His
 20 435 440 445
 Tyr Asp Asn Thr Gly Lys Asn Leu Lys Thr Gln Val Ile Gln Glu Asn
 450 455 460
 Val Asp Pro Val Thr Asn Arg Asp Tyr Ser Ile Phe Gly Trp Asn Asn
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 35 40 45
 Ser Asp Ser Ala Thr Val Lys Glu Thr Ser Ser Asn Met Gln Ser Pro
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 Gln Asn Ala Thr Ala Asn Gln Ser Thr Thr Lys Thr Ser Asn Val Thr
 65 70 75 80
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 85 90 95
 Asn Leu Thr Gln Ala Lys Asp Val Ser Thr Thr Pro Lys Thr Thr Thr
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 Ile Lys Pro Arg Thr Leu Asn Arg Met Ala Val Asn Thr Val Ala Ala
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 Pro Gln Gln Gly Thr Asn Val Asn Asp Lys Val His Phe Ser Asn Ile
 130 135 140

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Asp Ile Ala Ile Asp Lys Gly His Val Asn Gln Thr Thr Gly Lys Thr
 145 150 155 160
 5 Glu Phe Trp Ala Thr Ser Ser Asp Val Leu Lys Leu Lys Ala Asn Tyr
 165 170 175
 Thr Ile Asp Asp Ser Val Lys Glu Gly Asp Thr Phe Thr Phe Lys Tyr
 180 185 190
 10 Gly Gln Tyr Phe Arg Pro Gly Ser Val Arg Leu Pro Ser Gln Thr Gln
 195 200 205
 Asn Leu Tyr Asn Ala Gln Gly Asn Ile Ile Ala Lys Gly Ile Tyr Asp
 210 215 220
 15 Ser Thr Thr Asn Thr Thr Thr Tyr Thr Phe Thr Asn Tyr Val Asp Gln
 225 230 235 240
 Tyr Thr Asn Val Arg Gly Ser Phe Glu Gln Val Ala Phe Ala Lys Arg
 245 250 255
 20 Lys Asn Ala Thr Thr Asp Lys Thr Ala Tyr Lys Met Glu Val Thr Leu
 260 265 270
 Gly Asn Asp Thr Tyr Ser Glu Glu Ile Ile Val Asp Tyr Gly Asn Lys
 275 280 285
 25 Lys Ala Gln Pro Leu Ile Ser Ser Thr Asn Tyr Ile Asn Asn Glu Asp
 290 295 300
 Leu Ser Arg Asn Met Thr Ala Tyr Val Asn Gln Pro Lys Asn Thr Tyr
 305 310 315 320
 Thr Lys Gln Thr Phe Val Thr Asn Leu Thr Gly Tyr Lys Phe Asn Pro
 325 330 335
 30 Asn Ala Lys Asn Phe Lys Ile Tyr Glu Val Thr Asp Gln Asn Gln Phe
 340 345 350
 Val Asp Ser Phe Thr Pro Asp Thr Ser Lys Leu Lys Asp Val Thr Asp
 355 360 365
 35 Gln Phe Asp Val Ile Tyr Ser Asn Asp Asn Lys Thr Ala Thr Val Asp
 370 375 380
 Leu Met Lys Gly Gln Thr Ser Ser Asn Lys Gln Tyr Ile Ile Gln Gln
 385 390 395 400
 40 Val Ala Tyr Pro Asp Asn Ser Ser Thr Asp Asn Gly Lys Ile Asp Tyr
 405 410 415
 Thr Leu Asp Thr Asp Lys Thr Lys Tyr Ser Trp Ser Asn Ser Tyr Ser
 420 425 430
 45 Asn Val Asn Gly Ser Ser Thr Ala Asn Gly Asp Gln Lys Lys Tyr Asn
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10 Asn His Asn Lys Gln Val His Asn Glu Ser Gln Thr Ser Glu Asp Asn
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Val Gln Asn Glu Ala Gly Thr Ile Asp Asp Arg Gln Val Glu Ser Ser
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15 His Ser Thr Glu Ser Gln Glu Pro Ser His Gln Asp Ser Thr Pro Gln
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His Glu Glu Glu Tyr Tyr Asn Lys Asn Ala Phe Ala Met Asp Lys Ser
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20 His Pro Glu Pro Ile Glu Asp Asn Asp Lys His Asp Thr Ile Lys Asn
130 135 140
Ala Glu Asn Asn Thr Glu His Ser Thr Val Ser Asp Lys Ser Glu Ala
145 150 155 160
25 Glu Gln Ser Gln Gln Pro Lys Pro Tyr Phe Thr Thr Gly Ala Asn Gln
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35 40 45
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50 Asn Ile Glu Thr Val Lys Glu Glu Val Val Lys Glu Glu Ala Lys Pro
85 90 95
Gln Val Lys Glu Thr Thr Gln Ser Gln Asp Asn Ser Gly Asp Gln Arg
100 105 110
55 Gln Val Asp Leu Thr Pro Lys Lys Ala Thr Gln Asn Gln Val Ala Glu
115 120 125
Thr Gln Val Glu Val Ala Gln Pro Arg Thr Ala Ser Glu Ser Lys Pro
130 135 140

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Arg Val Thr Arg Ser Ala Asp Val Ala Glu Ala Lys Glu Ala Ser Asn
 145 150 155 160
 5 Ala Lys Val Glu Thr Gly Thr Asp Val Thr Ser Lys Val Thr Val Glu
 165 170 175
 Ile Gly Ser Ile Glu Gly His Asn Asn Thr Asn Lys Val Glu Pro His
 180 185 190
 10 Ala Gly Gln Arg Ala Val Leu Lys Tyr Lys Leu Lys Phe Glu Asn Gly
 195 200 205
 Leu His Gln Gly Asp Tyr Phe Asp Phe Thr Leu Ser Asn Asn Val Asn
 210 215 220
 15 Thr His Gly Val Ser Thr Ala Arg Lys Val Pro Glu Ile Lys Asn Gly
 225 230 235 240
 Ser Val Val Met Ala Thr Gly Glu Val Leu Glu Gly Gly Lys Ile Arg
 245 250 255
 20 Tyr Thr Phe Thr Asn Asp Ile Glu Asp Lys Val Asp Val Thr Ala Glu
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 275 280 285
 25 Asn Gln Thr Ile Thr Ser Thr Leu Asn Glu Glu Gln Thr Ser Lys Glu
 290 295 300
 Leu Asp Val Lys Tyr Lys Asp Gly Ile Gly Asn Tyr Tyr Ala Asn Leu
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 355 360 365
 Val Arg Ile Phe Glu Tyr Leu Gly Asn Asn Glu Asp Ile Ala Lys Ser
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 35 40 45
 Thr Ala Val Asp Ser Glu Ala Gly His Val Gly Gly Tyr Thr Glu Ser
 50 55 60
 15 Ser Glu Glu Ser Asn Pro Ile Asp Phe Glu Glu Ser Thr His Glu Asn
 65 70 75 80
 Ser Lys His His Ala Asp Val Val Glu Tyr Glu Glu Asp Thr Asn Pro
 85 90 95
 20 Gly Gly Gly Gln Val Thr Thr Glu Ser Asn Leu Val Glu Phe Asp Glu
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 Glu Ser Thr Lys Gly Ile Val Thr Gly Ala Val Ser Asp His Thr Thr
 115 120 125
 25 Val Glu Asp Thr Lys Glu Tyr Thr Thr Glu Ser Asn Leu Ile Glu Leu
 130 135 140
 Val Asp Glu Leu Pro Glu Glu His Gly Gln Ala Gln Gly Pro Val Glu
 145 150 155 160
 30 Glu Ile Thr Lys Asn Asn His His Ile Ser His Ser Gly Leu Gly Thr
 165 170 175
 Glu Asn Gly His Gly Asn Tyr Asp Val Ile Glu Glu Ile Glu Glu Asn
 180 185 190
 35 Ser His Val Asp Ile Lys Ser Glu Leu Gly Tyr Glu Gly Gly Gln Asn
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 Ser Gly Asn Gln Ser Phe Glu Glu Asp Thr Glu Glu Asp Lys Pro Lys
 210 215 220
 40 Tyr Glu Gln Gly Gly Asn Ile Val Asp Ile Asp Phe Asp Ser Val Pro
 225 230 235 240
 Gln Ile His Gly Gln Asn Lys Gly Asn Gln Ser Phe Glu Glu Asp Thr
 245 250 255
 45 Glu Lys Asp Lys Pro Lys Tyr Glu His Gly Gly Asn Ile Ile Asp Ile
 260 265 270
 Asp Phe Asp Ser Val Pro His Ile His Gly Phe Asn Lys His Thr Glu
 275 280 285
 50 Ile Ile Glu Glu Asp Thr Asn Lys Asp Lys Pro Ser Tyr Gln Phe Gly
 290 295 300
 Gly His Asn Ser Val Asp Phe Glu Glu Asp Thr Leu Pro Lys Val Ser
 305 310 315 320
 55 Gly Gln Asn Glu Gly Gln Gln Thr Ile Glu Glu Asp Thr Thr Pro Pro
 325 330 335
 Ile Val Pro Pro Thr Pro Pro Thr Pro Glu Val Pro Ser Glu Pro Glu

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10	Lys	Glu	Glu	Pro	Lys 405	Lys	Pro	Ser	Lys	Pro 410	Val	Glu	Gln	Gly	Lys 415	Val
	Val	Thr	Pro	Val 420	Ile	Glu	Ile	Asn	Glu 425	Lys	Val	Lys	Ala	Val 430	Ala	Pro
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	Ser	Val	Ile	Leu 100	Glu	Glu	Lys	Lys	Ala 105	Pro	Asn	Asn	Thr	Asn	Asn	Asp
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	Ser	Thr	Lys 195	Pro	Val	Ala	Thr	Ala 200	Pro	Thr	Ser	Val	Ala 205	Pro	Lys	Arg
55	Val	Asn 210	Ala	Lys	Met	Arg	Phe 215	Ala	Val	Ala	Gln	Pro 220	Ala	Ala	Val	Ala
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Val Gly Asp Gly Lys Asp Asn Val Ala Ala Ala His Asp Gly Lys Asp
 245 250 255
 5 Ile Glu Tyr Asp Thr Glu Phe Thr Ile Asp Asn Lys Val Lys Gly
 260 265 270
 Asp Thr Met Thr Ile Asn Tyr Asp Lys Asn Val Ile Pro Ser Asp Leu
 275 280 285
 10 Thr Asp Lys Asn Asp Pro Ile Asp Ile Thr Asp Pro Ser Gly Glu Val
 290 295 300
 Ile Ala Lys Gly Thr Phe Asp Lys Ala Thr Lys Gln Ile Thr Tyr Thr
 305 310 315 320
 15 Phe Thr Asp Tyr Val Asp Lys Tyr Glu Asp Ile Lys Ser Arg Leu Thr
 325 330 335
 Leu Tyr Ser Tyr Ile Asp Lys Lys Thr Val Pro Asn Glu Thr Ser Leu
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 20 Asn Leu Thr Phe Ala Thr Ala Gly Lys Glu Thr Ser Gln Asn Val Thr
 355 360 365
 Val Asp Tyr Gln Asp Pro Met Val His Gly Asp Ser Asn Ile Gln Ser
 370 375 380
 25 Ile Phe Thr Lys Leu Asp Glu Asp Lys Gln Thr Ile Glu Gln Gln Ile
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 405 410 415
 30 Ala Gly Ser Gln Val Asp Asp Tyr Gly Asn Ile Lys Leu Gly Asn Gly
 420 425 430
 Ser Thr Ile Ile Asp Gln Asn Thr Glu Ile Lys Val Tyr Lys Val Asn
 435 440 445
 35 Ser Asp Gln Gln Leu Pro Gln Ser Asn Arg Ile Tyr Asp Phe Ser Gln
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 Tyr Glu Asp Val Thr Ser Gln Phe Asp Asn Lys Lys Ser Phe Ser Asn
 465 470 475 480
 40 Asn Val Ala Thr Leu Asp Phe Gly Asp Ile Asn Ser Ala Tyr Ile Ile
 485 490 495
 Lys Val Val Ser Lys Tyr Thr Pro Thr Ser Asp Gly Glu Leu Asp Ile
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 45 Ala Gln Gly Thr Ser Met Arg Thr Thr Asp Lys Tyr Gly Tyr Tyr Asn
 515 520 525
 Tyr Ala Gly Tyr Ser Asn Phe Ile Val Thr Ser Asn Asp Thr Gly Gly
 530 535 540
 50 Gly Asp Gly Thr Val Lys Pro Glu Glu Lys Leu Tyr Lys Ile Gly Asp
 545 550 555 560
 Tyr Val Trp Glu Asp Val Asp Lys Asp Gly Val Gln Gly Thr Asp Ser
 565 570 575
 55 Lys Glu Lys Pro Ala Ser Gly Gly Gly Ser Ala Glu Ser Thr Asn Lys
 580 585 590
 Glu Leu Asn Glu Ala Thr Thr Ser Ala Ser Asp Asn Gln Ser Ser Asp
 595 600 605

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 5 Asn Gln Lys Glu Met Val Ser Ser Gln Gly Asn Glu Thr Thr Ser Asn
 625 630 635 640
 Gly Asn Lys Leu Ile Glu Lys Glu Ser Val Gln Ser Thr Thr Gly Asn
 645 650 655
 10 Lys Val Glu Val Ser Thr Ala Lys Ser Asp Glu Gln Ala Ser Pro Lys
 660 665 670
 Ser Thr Asn Glu Asp Leu Asn Thr Lys Gln Thr Ile Ser Asn Gln Glu
 675 680 685
 15 Ala Leu Gln Pro Asp Leu Gln Glu Asn Lys Ser Val Val Asn Val Gln
 690 695 700
 Pro Thr Asn Glu Glu Asn Lys Lys Val Asp Ala Lys Thr Glu Ser Thr
 705 710 715 720
 20 Thr Leu Asn Val Lys Ser Asp Ala Ile Lys Ser Asn Asp Glu Thr Leu
 725 730 735
 Val Asp Asn Asn Ser Asn Ser Asn Asn Glu Asn Asn Ala Asp Ile Ile
 740 745 750
 25 Leu Pro Lys Ser Thr Ala Pro Lys Arg Leu Asn Thr Arg Met Arg Ile
 755 760 765
 Ala Ala Val Gln Pro Ser Ser Thr Glu Ala Lys Asn Val Asn Asp Leu
 770 775 780
 30 Ile Thr Ser Asn Thr Thr Leu Thr Val Val Asp Ala Asp Lys Asn Asn
 785 790 795 800
 Lys Ile Val Pro Ala Gln Asp Tyr Leu Ser Leu Lys Ser Gln Ile Thr
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 35 Val Asp Asp Lys Val Lys Ser Gly Asp Tyr Phe Thr Ile Lys Tyr Ser
 820 825 830
 Asp Thr Val Gln Val Tyr Gly Leu Asn Pro Glu Asp Ile Lys Asn Ile
 835 840 845
 40 Gly Asp Ile Lys Asp Pro Asn Asn Gly Glu Thr Ile Ala Thr Ala Lys
 850 855 860
 His Asp Thr Ala Asn Asn Leu Ile Thr Tyr Thr Phe Thr Asp Tyr Val
 865 870 875 880
 45 Asp Arg Phe Asn Ser Val Gln Met Gly Ile Asn Tyr Ser Ile Tyr Met
 885 890 895
 Asp Ala Asp Thr Ile Pro Val Ser Lys Asn Asp Val Glu Phe Asn Val
 900 905 910
 50 Thr Ile Gly Asn Thr Thr Thr Lys Thr Thr Ala Asn Ile Gln Tyr Pro
 915 920 925
 Asp Tyr Val Val Asn Glu Lys Asn Ser Ile Gly Ser Ala Phe Thr Glu
 930 935 940
 55 Thr Val Ser His Val Gly Asn Lys Glu Asn Pro Gly Tyr Tyr Lys Gln
 945 950 955 960
 Thr Ile Tyr Val Asn Pro Ser Glu Asn Ser Leu Thr Asn Ala Lys Leu
 965 970 975
 Lys Val Gln Ala Tyr His Ser Ser Tyr Pro Asn Asn Ile Gly Gln Ile

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	980		985		990	
	Asn Lys Asp Val Thr Asp Ile Lys Ile Tyr Gln Val Pro Lys Gly Tyr					
	995		1000		1005	
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	1010		1015		1020	
	Thr Asn Gln Tyr Leu Gln Lys Ile Thr Tyr Gly Asp Asn Asn Ser Ala					
	1025		1030		1035	1040
10	Val Ile Asp Phe Gly Asn Ala Asp Ser Ala Tyr Val Val Met Val Asn					
		1045		1050		1055
	Thr Lys Phe Gln Tyr Thr Asn Ser Glu Ser Pro Thr Leu Val Gln Met					
		1060		1065		1070
15	Ala Thr Leu Ser Ser Thr Gly Asn Lys Ser Val Ser Thr Gly Asn Ala					
		1075		1080		1085
	Leu Gly Phe Thr Asn Asn Gln Ser Gly Gly Ala Gly Gln Glu Val Tyr					
		1090		1095		1100
20	Lys Ile Gly Asn Tyr Val Trp Glu Asp Thr Asn Lys Asn Gly Val Gln					
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	Glu Leu Gly Glu Lys Gly					
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	gcgaactggg aaggtaagc ttccagccgt ttcgaagagc aattccaaca acttagtcct 180					
	aaagtagaaa aatttgaca attattagaa gaaattaaac aacaattgaa tagcactgct 240					
	gatgccgttc aagaacaaga ccaacaactt tctaataatt tcggtttgca agctagcggg 300					
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	ttactcatta tggcaaagt tcaagaagaa ttagtacaac cgatggctga ccatcaaaaa 540					
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	Thr Ala Lys Ala Glu Asn Ser Val Lys Leu Ile Thr Asn Thr Asn Val					
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	Val Gly Asn His Thr Ile Ile Thr Asn Lys His Val Thr Tyr His Met					
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 Glu Asp Ile Ala Val Val Gln Val Glu Glu Lys Ser Thr Gln Pro Lys
 115 125
 Gly Arg Lys Phe Lys Asp Phe Thr Ser Lys Phe Asn Ile Ala Ser Glu
 130 135 140
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 Ala Lys Glu Asn Glu Pro Ile Ser Val Ile Gly Tyr Pro Asn Pro Asn
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 Gly Asn Lys Leu Gln Met Tyr Glu Ser Thr Gly Lys Val Leu Ser Val
 165 170 175
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 Asn Gly Asn Ile Val Thr Ser Asp Ala Val Val Gln Pro Gly Ser Ser
 180 185 190
 Gly Ser Pro Ile Leu Asn Ser Lys Arg Glu Ala Ile Gly Val Met Tyr
 195 200 205
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<211> 203
<212> PRT
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35 40 45

40 Gln Lys Val Glu Glu Val Pro Asn Asn Ser Glu Lys Ala Leu Val Lys
50 60

Lys Leu Tyr Asp Arg Tyr Ser Lys Asp Thr Ile Asn Gly Lys Ser Asn
65 70 75 80

45 Lys Ser Arg Asn Trp Val Tyr Ser Glu Arg Pro Leu Asn Glu Asn Gln
85 90 95

Val Arg Ile His Leu Glu Gly Thr Tyr Thr Val Ala Gly Arg Val Tyr
100 105 110

50 Thr Pro Lys Arg Asn Ile Thr Leu Asn Lys Glu Val Val Thr Leu Lys
115 120 125

Glu Leu Asp His Ile Ile Arg Phe Ala His Ile Ser Tyr Gly Leu Tyr
130 135 140

55 Met Gly Glu His Leu Pro Lys Gly Asn Ile Val Ile Asn Thr Lys Asp
145 150 155 160

Gly Gly Lys Tyr Thr Leu Glu Ser His Lys Glu Leu Gln Lys Asp Arg

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165 170 175
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 35 40 45
 20 Thr Lys His Leu Thr Gly Thr Phe Ser Ser Lys Asn Gly Glu Thr Val
 50 55 60
 Glu Gly Lys Ala Glu Ile Lys Asn Gly Lys Leu Met Leu Thr Asn Tyr
 65 70 75 80
 25 Lys Ser Ser Lys Gly Pro Asp Leu Tyr Val Tyr Leu Thr Lys Asn Gly
 85 90 95
 Asp Ile Lys Asn Gly Lys Glu Ile Ala Met Val Asp Tyr Asp Lys Glu
 100 105 110
 30 Lys Gln Thr Phe Asp Leu Lys Asn Val Asp Leu Ser Lys Tyr Asp Glu
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 35 Leu Lys
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 35 40 45
 Asp Glu Thr Ser Lys Asp Thr Thr Ser Lys Asp Ile Asp Lys Ala Asp
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 Ile Asp Asp Ser Thr Ser Asp Ser Asn Asn Ile Ile Asp Phe Ile Tyr
 85 90 95
 55 Lys Asn Leu Pro Gln Thr Asn Ile Asn Gln Leu Leu Thr Lys Asn Lys
 100 105 110

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 5 Leu Asn Ser Asp Ile Ser Asp Tyr Glu Gln Pro Arg Asn Gly Glu Lys
 130 135 140
 Ser Thr Asn Asp Ser Asn Lys Asn Ser Asp Asn Ser Ile Lys Asn Asp
 145 150 155 160
 10 Thr Asp Thr Gln Ser Ser Lys Gln Asp Lys Ala Asp Asn Gln Lys Ala
 165 170 175
 Pro Lys Ser Asn Asn Thr Lys Pro Ser Thr Ser Asn Lys Gln Pro Asn
 180 185 190
 15 Ser Pro Lys Pro Thr Gln Pro Asn Gln Ser Asn Ser Gln Pro Ala Ser
 195 200 205
 Asp Asp Lys Ala Asn Gln Lys Ser Ser Ser Lys Asp Asn Gln Ser Met
 210 215 220
 20 Ser Asp Ser Ala Leu Asp Ser Ile Leu Asp Gln Tyr Ser Glu Asp Ala
 225 230 235 240
 Lys Lys Thr Gln Lys Asp Tyr Ala Ser Gln Ser Lys Lys Asp Lys Asn
 245 250 255
 25 Glu Lys Ser Asn Thr Lys Asn Pro Gln Leu Pro Thr Gln Asp Glu Leu
 260 265 270
 Lys His Lys Ser Lys Pro Ala Gln Ser Phe Asn Asn Asp Val Asn Gln
 275 280 285
 30 Lys Asp Thr Arg Ala Thr Ser Leu Phe Glu Thr Asp Pro Ser Ile Ser
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 Asn Asn Asp Asp Ser Gly Gln Phe Asn Val Val Asp Ser Lys Asp Thr
 305 310 315 320
 35 Arg Gln Phe Val Lys Ser Ile Ala Lys Asp Ala His Arg Ile Gly Gln
 325 330 335
 Asp Asn Asp Ile Tyr Ala Ser Val Met Ile Ala Gln Ala Ile Leu Glu
 340 345 350
 40 Ser Asp Ser Gly Arg Ser Ala Leu Ala Lys Ser Pro Asn His Asn Leu
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 Phe Gly Ile Lys Gly Ala Phe Glu Gly Asn Ser Val Pro Phe Asn Thr
 370 375 380
 45 Leu Glu Ala Asp Gly Asn Gln Leu Tyr Ser Ile Asn Ala Gly Phe Arg
 385 390 395 400
 Lys Tyr Pro Ser Thr Lys Glu Ser Leu Lys Asp Tyr Ser Asp Leu Ile
 405 410 415
 50 Lys Asn Gly Ile Asp Gly Asn Arg Thr Ile Tyr Lys Pro Thr Trp Lys
 420 425 430
 Ser Glu Ala Asp Ser Tyr Lys Asp Ala Thr Ser His Leu Ser Lys Thr
 435 440 445
 Tyr Ala Thr Asp Pro Asn Tyr Ala Lys Lys Leu Asn Ser Ile Ile Lys
 450 455 460
 55 His Tyr Gln Leu Thr Gln Phe Asp Asp Glu Arg Met Pro Asp Leu Asp
 465 470 475 480

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Lys Tyr Glu Arg Ser Ile Lys Asp Tyr Asp Asp Ser Ser Asp Glu Phe
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 Lys Pro Phe Arg Glu Val Ser Asp Ser Met Pro Tyr Pro His Gly Gln
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 5 Cys Thr Trp Tyr Val Tyr Asn Arg Met Lys Gln Phe Gly Thr Ser Ile
 515 520 525
 Ser Gly Asp Leu Gly Asp Ala His Asn Trp Asn Asn Arg Ala Gln Tyr
 530 535 540
 10 Arg Asp Tyr Gln Val Ser His Thr Pro Lys Arg His Ala Ala Val Val
 545 550 555 560
 Phe Glu Ala Gly Gln Phe Gly Ala Asp Gln His Tyr Gly His Val Ala
 565 570 575
 15 Phe Val Glu Lys Val Asn Ser Asp Gly Ser Ile Val Ile Ser Glu Ser
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 Asn Val Lys Gly Leu Gly Ile Ile Ser His Arg Thr Ile Asn Ala Ala
 595 600 605
 20 Ala Ala Glu Glu Leu Ser Tyr Ile Thr Gly Lys
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 <211> 208
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 25 <213> Staphylococcus aureus
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 Gln Val Asp Lys Ile Lys Asp Asp Glu Glu Pro Ile Lys Thr Val Gly
 35 40 45
 Lys Lys Ile Ala Glu Leu Asp Glu Lys Lys Lys Lys Leu Thr Glu Asp
 50 55 60
 Val Asn Ser Lys Asp Thr Ala Val Arg Gly Lys Ala Val Lys Asp Leu
 65 70 75 80
 40 Ile Lys Asn Ala Asp Asp Arg Leu Lys Glu Phe Glu Lys Glu Glu Asp
 85 90 95
 Ala Ile Lys Lys Ser Glu Gln Asp Phe Lys Lys Ala Lys Ser His Val
 100 105 110
 45 Asp Asn Ile Asp Asn Asp Val Lys Arg Lys Glu Val Lys Gln Leu Asp
 115 120 125
 Asp Val Leu Lys Glu Lys Tyr Lys Leu His Ser Asp Tyr Ala Lys Ala
 130 135 140
 50 Tyr Lys Lys Ala Val Asn Ser Glu Lys Thr Leu Phe Lys Tyr Leu Asn
 145 150 155 160
 Gln Asn Asp Ala Thr Gln Gln Gly Val Asn Glu Lys Ser Lys Ala Ile
 165 170 175
 55 Glu Gln Asn Tyr Lys Lys Leu Lys Glu Val Ser Asp Lys Tyr Thr Lys
 180 185 190
 Val Leu Asn Lys Val Gly Lys Glu Lys Gln Asp Val Asp Gln Phe Lys

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	Gln Ile Lys Gly Ala Asn Asp Ile Val Ile Lys Lys Gly Gln Asp Tyr			
	35 40 45			
15	Asn Leu Leu Asn Gly Ile Ser Ala Phe Asp Lys Glu Asp Gly Asp Leu			
	50 55 60			
	Thr Asp Lys Ile Lys Val Asp Gly Gln Ile Asp Thr Ser Lys Ser Gly			
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	20 25 30			
35	Leu Lys Val Val Thr Thr Asn Ser Ile Leu Tyr Asp Met Ala Lys Asn			
	35 40 45			
	Val Gly Gly Asp Asn Val Asp Ile His Ser Ile Val Pro Val Gly Gln			
	50 55 60			
40	Asp Pro His Glu Tyr Glu Val Lys Pro Lys Asp Ile Lys Lys Leu Thr			
	65 70 75 80			
	Asp Ala Asp Val Ile Leu Tyr Asn Gly Leu Asn Leu Glu Thr Gly Asn			
	85 90 95			
45	Gly Trp Phe Glu Lys Ala Leu Glu Gln Ala Gly Lys Ser Leu Lys Asp			
	100 105 110			
	Lys Lys Val Ile Ala Val Ser Lys Asp Val Lys Pro Ile Tyr Leu Asn			
	115 120 125			
50	Gly Glu Glu Gly Asn Lys Asp Lys Gln Asp Pro His Ala Trp Leu Ser			
	130 135 140			
	Leu Asp Asn Gly Ile Lys Tyr Val Lys Thr Ile Gln Gln Thr Phe Ile			
	145 150 155 160			
	Asp Asn Asp Lys Lys His Lys Ala Asp Tyr Glu Lys Gln Gly Asn Lys			
	165 170 175			
55	Tyr Ile Ala Gln Leu Glu Lys Leu Asn Asn Asp Ser Lys Asp Ser Lys			
	180 185 190			

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Asp Lys Phe Asn Asp Ile Pro Lys Glu Gln Arg Ala Met Ile Thr Ser
 195 200 205
 5 Glu Gly Ala Phe Lys Tyr Phe Ser Lys Gln Tyr Gly Ile Thr Pro Gly
 210 215 220
 Tyr Ile Trp Glu Ile Asn Thr Glu Lys Gln Gly Thr Pro Glu Gln Met
 225 230 235 240
 10 Arg Gln Ala Ile Glu Phe Val Lys Lys His Lys Leu Lys His Leu Leu
 245 250 255
 Val Glu Thr Ser Val Asp Lys Lys Ala Met Glu Ser Leu Ser Glu Glu
 260 265 270
 15 Thr Lys Lys Asp Ile Phe Gly Glu Val Tyr Thr Asp Ser Ile Gly Lys
 275 280 285
 Glu Gly Thr Lys Gly Asp Ser Tyr Tyr Lys Met Met Lys Ser Asn Ile
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 20 Glu Thr Val His Gly Ser Met Lys
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 Phe Ser Leu Gly Val Lys Gly Leu Val Gln Asn Leu Ile Leu Leu Met
 35 40 45
 Asn Pro Tyr Ser Leu Val Ala Leu Val Leu Ser Val Phe Leu Phe Phe
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 35 Lys Gly Lys Lys Ala Phe Trp Phe Met Phe Ile Gly Gly Phe Leu Leu
 65 70 75 80
 40 Thr Phe Leu Leu Tyr Ala Asn Val Val Tyr Phe Arg Phe Phe Ser Asp
 85 90 95
 45 Phe Leu Thr Phe Ser Thr Leu Asn Gln Val Gly Asn Val Glu Ser Met
 100 105 110
 Gly Gly Ala Val Ser Ala Ser Phe Lys Trp Tyr Asp Phe Val Tyr Phe
 115 120 125
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 130 135 140
 Leu Asp Thr Lys Ala Phe Ser Lys Lys Phe Val Pro Val Val Met Ala
 145 150 155 160
 55 Ala Ser Val Ala Leu Phe Phe Leu Asn Leu Ala Phe Ala Glu Thr Asp
 165 170 175
 Arg Pro Glu Leu Leu Thr Arg Thr Phe Asp His Lys Tyr Leu Val Lys
 180 185 190
 55 Tyr Leu Gly Pro Tyr Asn Phe Thr Val Tyr Asp Gly Val Lys Thr Ile
 195 200 205

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5
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 15
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 55

Glu Asn Asn Gln Gln Lys Ala Leu Ala Ser Glu Asp Asp Leu Thr Lys
 210 215 220
 Val Leu Asn Tyr Thr Lys Gln Arg Gln Thr Glu Pro Asn Pro Glu Tyr
 225 230 235 240
 Tyr Gly Val Ala Lys Lys Lys Asn Ile Ile Lys Ile His Leu Glu Ser
 245 250 255
 Phe Gln Thr Phe Leu Ile Asn Lys Lys Val Asn Gly Lys Glu Val Thr
 260 265 270
 Pro Phe Leu Asn Lys Leu Ser Ser Gly Lys Glu Gln Phe Thr Tyr Phe
 275 280
 Pro Asn Phe Phe His Gln Thr Gly Gln Gly Lys Thr Ser Asp Ser Glu
 290 295 300
 Phe Thr Met Asp Asn Ser Leu Tyr Gly Leu Pro Gln Gly Ser Ala Phe
 305 310 315 320
 Ser Leu Lys Gly Asp Asn Thr Tyr Gln Ser Leu Pro Ala Ile Leu Asp
 325 330 335
 Gln Lys Gln Gly Tyr Lys Ser Asp Val Met His Gly Asp Tyr Lys Thr
 340 345 350
 Phe Trp Asn Arg Asp Gln Val Tyr Lys His Phe Gly Ile Asp Lys Phe
 355 360 365
 Tyr Asp Ala Thr Tyr Tyr Asp Met Ser Asp Lys Asn Val Val Asn Leu
 370 375 380
 Gly Leu Lys Asp Lys Ile Phe Phe Lys Asp Ser Ala Asn Tyr Gln Ala
 385 390 395 400
 Lys Met Lys Ser Pro Phe Tyr Ser His Leu Ile Thr Leu Thr Asn His
 405 410 415
 Tyr Pro Phe Thr Leu Asp Glu Lys Asp Ala Thr Ile Glu Lys Ser Asn
 420 425 430
 Thr Gly Asp Ala Thr Val Asp Gly Tyr Ile Gln Thr Ala Arg Tyr Leu
 435 440 445
 Asp Glu Ala Leu Glu Glu Tyr Ile Asn Asp Leu Lys Lys Lys Gly Leu
 450 455 460
 Tyr Asp Asn Ser Val Ile Met Ile Tyr Gly Asp His Tyr Gly Ile Ser
 465 470 475 480
 Glu Asn His Asn Asn Ala Met Glu Lys Leu Leu Gly Glu Lys Ile Thr
 485 490 495
 Pro Ala Lys Phe Thr Asp Leu Asn Arg Thr Gly Phe Trp Ile Lys Ile
 500 505 510
 Pro Gly Lys Ser Gly Gly Ile Asn Asn Glu Tyr Ala Gly Gln Val Asp
 515 520 525
 Val Met Pro Thr Ile Leu His Leu Ala Gly Ile Asp Thr Lys Asn Tyr
 530 535 540
 Leu Met Phe Gly Thr Asp Leu Phe Ser Lys Gly His Asn Gln Val Val
 545 550 555 560
 Pro Phe Arg Asn Gly Asp Phe Ile Thr Lys Asp Tyr Lys Tyr Val Asn
 565 570 575
 Gly Lys Ile Tyr Ser Asn Lys Asn Asn Glu Leu Ile Thr Thr Gln Pro

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580 585 590

Ala Asp Phe Glu Lys Asn Lys Lys Gln Val Glu Lys Asp Leu Glu Met
595 600 605

5 Ser Asp Asn Val Leu Asn Gly Asp Leu Phe Arg Phe Tyr Lys Asn Pro
610 615 620

Asp Phe Lys Lys Val Asn Pro Ser Lys Tyr Lys Tyr Glu Thr Gly Pro
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Ile Ser Phe Leu Pro Ala Leu Gly Phe Asn Gln Ile Leu Leu Glu Arg
35 40 45

25 Glu Glu Asp Gln Leu Asn Ile Met Asn Ser Ala Thr Glu Glu His His
50 55 60

His Lys Asp Tyr Ile Lys Leu Tyr Asn Leu Gly Gly Gly Ala Ala Lys
65 70 75 80

30 Lys Ile Ala Ile Glu Val Leu Leu Gly Lys Asp Lys Val Ile Gln Lys
85 90 95

Lys Tyr Val His Ile Leu Pro Ser Lys Glu Gly Tyr Met Leu Pro Ile
100 105 110

35 Asn Lys Asn Val Tyr Glu Glu Leu Glu Arg Thr Ile Glu Asn Asn Gly
115 120 125

His Glu Ala Asp Leu Asn Val Arg Met Thr Tyr Tyr His Asn Val Ser
130 135 140

40 Arg Lys Gln Gln Glu Val Ile Leu Lys Gly Gln Ile Asp Arg Phe Asn
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Thr Tyr Asn Asn Lys Glu Ile Tyr Asp Leu Gln Phe Ile
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35 40 45

55 Ile Thr Val Glu His Ala Gln Ile Asn Ile Phe Gln Ser Asn Ser Asn
50 55 60

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Ser Asn Leu Met Glu Phe Asn Ile Leu Thr Met Gly Gly Lys Ser Gly
 65 70 75 80
 5 Ala Met Val Gly Tyr Ser Glu Ile Asp Ser Ser His Phe Thr Asp Arg
 85 90
 Asp Lys Arg Val Ile Arg Arg Asp His Val Lys Glu Ala Gln Ser Leu
 100 105 110
 10 Val Glu Asn Tyr Lys Asp Thr Gln Ser Ala Asp Ala Arg Met Lys Ala
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 130 135 140
 15 Lys Gln Ile Asp Lys Val Tyr Asn Gly Leu Gln Arg
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 35 40 45
 Lys Asp Asp Glu Arg Ile Asp Lys Leu Met Lys Glu Ser Asp Lys Asn
 50 55 60
 30 Leu Thr Pro Tyr Lys Ala Glu Thr Val Asn Asp Leu Tyr Leu Ile Val
 65 70 75 80
 Lys Lys Leu Ser Gln Gly Asp Val Lys Lys Ala Val Val Arg Ile Lys
 85 90 95
 35 Asp Gly Gly Pro Arg Asp Tyr Tyr Thr Phe Asp Leu Thr Arg Pro Leu
 100 105 110
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 115 120 125
 40 Ser Ile Thr Trp Tyr
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 45 <210> 187
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 Asp Asn Gly Ala Ser Asp Lys Leu Leu Ser Asn Val Lys Lys Glu Tyr
 35 40 45
 55 Pro Tyr Gln Thr Pro Val Leu Gly Arg Ser Gln Ser Gly Trp Asp Lys
 50 55 60

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Thr Glu Gly Ser Tyr Ser Ser Thr Val Ala Glu Asp Gly Gly Val Ala
 65 70 75 80
 Ile Val Ser Lys Tyr Pro Ile Lys Glu Lys Ile Gln His Val Phe Lys
 5 85 90 95
 Ser Gly Cys Gly Phe Asp Asn Asp Ser Asn Lys Gly Phe Val Tyr Thr
 100 105 110
 Lys Ile Glu Lys Asn Gly Lys Asn Val His Val Ile Gly Thr His Thr
 115 120 125
 10 Gln Ser Glu Asp Ser Arg Cys Gly Ala Gly His Asp Arg Lys Ile Arg
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 Ala Glu Gln Met Lys Glu Ile Ser Asp Phe Val Lys Lys Lys Asn Ile
 145 150 155 160
 15 Pro Lys Asp Glu Thr Val Tyr Ile Gly Gly Asp Leu Asn Val Asn Lys
 165 170 175
 Gly Thr Pro Glu Phe Lys Asp Met Leu Lys Asn Leu Asn Val Asn Asp
 180 185 190
 20 Val Leu Tyr Ala Gly His Asn Ser Thr Trp Asp Pro Gln Ser Asn Ser
 195 200 205
 Ile Ala Lys Tyr Asn Tyr Pro Asn Gly Lys Pro Glu His Leu Asp Tyr
 210 215 220
 25 Ile Phe Thr Asp Lys Asp His Lys Gln Pro Lys Gln Leu Val Asn Glu
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 35 40 45
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 5 Pro Ser Ala Phe Lys Val Gln Leu Gln Leu Pro Asp Asn Glu Val Ala
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 55 Val Asn His His Phe Ile Thr Thr Gln Thr His Tyr Lys Lys Val Ile
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Pro Gly Tyr His Ala His Lys Phe Val Thr Pro Gly His Ala Ser Ile
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 5 Lys Ile Asn His Phe Cys Val Val Pro Gln Ile Asn Ser Phe Lys Val
 195 200 205
 Ile Pro Pro Tyr Gly His Asn Ser His Arg Met His Val Pro Ser Phe
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 10 Gln Asn Asn Thr Thr Ala Thr His Gln Asn Ala Lys Val Asn Lys Ala
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 15 Lys Tyr Phe Ser Phe Ser Gln Ser Asn Gly Tyr Lys Ile Gly Lys Pro
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 210 215 220
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 225 230 235 240
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 15 Leu Thr Lys Thr Tyr Thr Phe Val Phe Thr Asp Tyr Val Asn Asn Lys
 260 265 270
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 290 295 300
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 25 Gly Ile Asp Lys Pro Asn Gly Ala Asn Ile Ser Ser Gln Ile Ile Gly
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 405 410 415
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 325 330 335
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		290					295					300					
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	Ser	Lys	Leu	Gly	Asn	Ile	Val	Pro	Glu	Tyr	Lys	Glu	Ile	Asn	Asn	Arg	
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25	Val	Asn	Val	Ala	Thr	Asn	Asn	Pro	Ala	Ser	Gln	Gln	Val	Asp	Lys	His	
			355					360					365				
	Phe	Val	Ala	Lys	Gly	Pro	Glu	Val	Asn	Arg	Phe	Ile	Thr	Gln	Asn	Lys	
		370					375					380					
30	Val	Asn	His	His	Phe	Ile	Thr	Thr	Gln	Thr	His	Tyr	Lys	Lys	Val	Ile	
					390						395					400	
	Thr	Ser	Tyr	Lys	Ser	Thr	His	Val	His	Lys	His	Val	Asn	His	Ala	Lys	
				405						410					415		
35	Asp	Ser	Ile	Asn	Lys	His	Phe	Ile	Val	Lys	Pro	Ser	Glu	Ser	Pro	Arg	
				420					425					430			
	Tyr	Thr	His	Pro	Ser	Gln	Ser	Leu	Ile	Ile	Lys	His	His	Phe	Ala	Val	
			435					440					445				
40	Pro	Gly	Tyr	His	Ala	His	Lys	Phe	Val	Thr	Pro	Gly	His	Ala	Ser	Ile	
		450					455					460					
	Lys	Ile	Asn	His	Phe	Cys	Val	Val	Pro	Gln	Ile	Asn	Ser	Phe	Lys	Val	
		465				470					475					480	
45	Ile	Pro	Pro	Tyr	Gly	His	Asn	Ser	His	Arg	Met	His	Val	Pro	Ser	Phe	
					485					490					495		
	Gln	Asn	Asn	Thr	Thr	Ala	Thr	His	Gln	Asn	Ala	Lys	Val	Asn	Lys	Ala	
				500					505					510			
50	Tyr	Asp	Tyr	Lys	Tyr	Phe	Tyr	Ser	Tyr	Lys	Val	Val	Lys	Gly	Val	Lys	
			515					520					525				
	Lys	Tyr	Phe	Ser	Phe	Ser	Gln	Ser	Asn	Gly	Tyr	Lys	Ile	Gly	Lys	Pro	
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	Ser	Pro	Thr	His	Tyr	Val	Pro	Glu	Phe	Lys	Gly	Ser	Leu	Pro	Ala	Pro	
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Arg Val

5
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 <212> PRT
 <213> Staphylococcus aureus

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 Gln Val Gln Lys Asp Gly Ser Ser Glu Lys Ser His Met Asp Asp Tyr
 15 35 40 45
 Met Gln His Pro Gly Lys Val Ile Lys Gln Asn Asn Lys Tyr Tyr Phe
 50 55 60
 Gln Thr Val Leu Asn Asn Ala Ser Phe Trp Lys Glu Tyr Lys Phe Tyr
 20 65 70 75 80
 Asn Ala Asn Asn Gln Glu Leu Ala Thr Thr Val Val Asn Asp Asn Lys
 85 90 95
 Lys Ala Asp Thr Arg Thr Ile Asn Val Ala Val Glu Pro Gly Tyr Lys
 100 105 110
 25 Ser Leu Thr Thr Lys Val His Ile Val Val Pro Gln Ile Asn Tyr Asn
 115 120 125
 His Arg Tyr Thr Thr His Leu Glu Phe Glu Lys Ala Ile Pro Thr Leu
 130 135 140
 30 Ala Ala Ser Gly Gly Gly Ser Met Ala Met Ile Lys Met Ser Pro Glu
 145 150 155 160
 Glu Ile Arg Ala Lys Ser Gln Ser Tyr Gly Gln Gly Ser Asp Gln Ile
 165 170 175
 35 Arg Gln Ile Leu Ser Asp Leu Thr Arg Ala Gln Gly Glu Ile Ala Ala
 180 185 190
 Asn Trp Glu Gly Gln Ala Phe Ser Arg Phe Glu Glu Gln Phe Gln Gln
 195 200 205
 40 Leu Ser Pro Lys Val Glu Lys Phe Ala Gln Leu Leu Glu Glu Ile Lys
 210 215 220
 Gln Gln Leu Asn Ser Thr Ala Asp Ala Val Gln Glu Gln Asp Gln Gln
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 45 Leu Ser Asn Asn Phe Gly Leu Gln
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 <212> PRT
 <213> Staphylococcus aureus

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 Asn Gln Ser Thr Gln Val Ser Gln Ala Thr Ser Gln Pro Ile Asn Phe
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Gln Val Gln Lys Asp Gly Ser Ser Glu Lys Ser His Met Asp Asp Tyr
 35 40 45
 Met Gln His Pro Gly Lys Val Ile Lys Gln Asn Asn Lys Tyr Tyr Phe
 5 50 60
 Gln Thr Val Leu Asn Asn Ala Ser Phe Trp Lys Glu Tyr Lys Phe Tyr
 65 70 75 80
 Asn Ala Asn Asn Gln Glu Leu Ala Thr Thr Val Val Asn Asp Asn Lys
 10 85 90 95
 Lys Ala Asp Thr Arg Thr Ile Asn Val Ala Val Glu Pro Gly Tyr Lys
 100 105
 Ser Leu Thr Thr Lys Val His Ile Val Val Pro Gln Ile Asn Tyr Asn
 115 120 125
 15 His Arg Tyr Thr Thr His Leu Glu Phe Glu Lys Ala Ile Pro Thr Leu
 130 135 140
 Ala Ala Ser Gly Gly Gly Ser Met Ala Met Ile Lys Met Ser Pro Glu
 145 150 155 160
 20 Glu Ile Arg Ala Lys Ser Gln Ser Tyr Gly Gln Gly Ser Asp Gln Ile
 165 170 175
 Arg Gln Ile Leu Ser Asp Leu Thr Arg Ala Gln Gly Glu Ile Ala Ala
 180 185 190
 25 Asn Trp Glu Gly Gln Ala Phe Ser Arg Phe Glu Glu Gln Phe Gln Gln
 195 200 205
 Leu Ser Pro Lys Val Glu Lys Phe Ala Gln Leu Leu Glu Glu Ile Lys
 210 215 220
 30 Gln Gln Leu Asn Ser Thr Ala Asp Ala Val Gln Glu Gln Asp Gln Gln
 225 230 235 240
 Leu Ser Asn Asn Phe Gly Leu Gln Ala Ser Gly Gly Gly Ser Met Gly
 245 250 255
 35 Gly Tyr Lys Gly Ile Lys Ala Asp Gly Gly Lys Val Asp Gln Ala Lys
 260 265 270
 Gln Leu Ala Ala Lys Thr Ala Lys Asp Ile Glu Ala Cys Gln Lys Gln
 275 280 285
 40 Thr Gln Gln Leu Ala Glu Tyr Ile Glu Gly Ser Asp Trp Glu Gly Gln
 290 295 300
 Phe Ala Asn Lys Val Lys Asp Val Leu Leu Ile Met Ala Lys Phe Gln
 305 310 315 320
 45 Glu Glu Leu Val Gln Pro Met Ala Asp His Gln Lys Ala Ile Asp Asn
 325 330 335
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 340 345 350
 50 Leu Asp Arg Val Asn Pro
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 <212> PRT
 <213> Staphylococcus aureus
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5	Gln	Thr	Val	Phe	Val	Asn	Pro	Lys	Gln	Arg	Val	Leu	Gly	Asn	Thr	Trp
			35					40					45			
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		50					55					60				
10	Val	Ser	Ala	Thr	Asp	Thr	Lys	Leu	Arg	Ile	Phe	Glu	Val	Asn	Asp	Thr
						70					75					80
	Ser	Lys	Leu	Ser	Asp	Ser	Tyr	Tyr	Ala	Asp	Pro	Asn	Asp	Ser	Asn	Leu
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15	Lys	Glu	Val	Thr	Asp	Gln	Phe	Lys	Asn	Arg	Ile	Tyr	Tyr	Glu	His	Pro
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	Asn	Val	Ala	Ser	Ile	Lys	Phe	Gly	Asp	Ile	Thr	Lys	Thr	Tyr	Val	Val
			115					120					125			
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	Val	Ile	Gln	Glu	Asn	Val	Asp	Pro	Val	Thr	Asn	Arg	Asp	Tyr	Ser	Ile
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25	Phe	Gly	Trp	Asn	Asn	Glu	Asn	Val	Val	Arg	Tyr	Gly	Gly	Gly	Ser	Ala
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	Asp	Gly	Asp	Ser	Ala	Val	Asn	Ala	Ser	Gly	Gly	Gly	Ser	Pro	Asp	Tyr
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30	Val	Val	Asn	Glu	Lys	Asn	Ser	Ile	Gly	Ser	Ala	Phe	Thr	Glu	Thr	Val
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	Ser	His	Val	Gly	Asn	Lys	Glu	Asn	Pro	Gly	Tyr	Tyr	Lys	Gln	Thr	Ile
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35	Tyr	Val	Asn	Pro	Ser	Glu	Asn	Ser	Leu	Thr	Asn	Ala	Lys	Leu	Lys	Val
						230					235					240
	Gln	Ala	Tyr	His	Ser	Ser	Tyr	Pro	Asn	Asn	Ile	Gly	Gln	Ile	Asn	Lys
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40	Asp	Val	Thr	Asp	Ile	Lys	Ile	Tyr	Gln	Val	Pro	Lys	Gly	Tyr	Thr	Leu
				260					265					270		
	Asn	Lys	Gly	Tyr	Asp	Val	Asn	Thr	Lys	Glu	Leu	Thr	Asp	Val	Thr	Asn
			275					280					285			
45	Gln	Tyr	Leu	Gln	Lys	Ile	Thr	Tyr	Gly	Asp	Asn	Asn	Ser	Ala	Val	Ile
		290					295					300				
	Asp	Phe	Gly	Asn	Ala	Asp	Ser	Ala	Tyr	Val	Val	Met	Val	Asn	Thr	Lys
						310					315					320
50	Phe	Gln	Tyr	Thr	Asn	Ser	Glu	Ser	Pro	Thr	Leu	Val	Gln	Met	Ala	Thr
					325					330					335	
	Leu	Ser	Ser	Thr	Gly	Asn	Lys	Ser	Val	Ser	Thr	Gly	Asn	Ala	Leu	Gly
				340					345					350		
55	Phe	Thr	Asn	Asn	Gln	Ser	Gly	Gly	Ala	Gly	Gln	Glu	Val	Tyr	Lys	Ile
			355				360						365			
	Gly	Asn	Tyr	Val	Trp	Glu	Asp	Thr	Asn	Lys	Asn	Gly	Val	Gln	Glu	Leu
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Gly Glu Lys Gly
385

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<210> 212
<211> 486
<212> PRT
<213> Staphylococcus aureus

<400> 212
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Gln Val Gln Lys Asp Gly Ser Ser Glu Lys Ser His Met Asp Asp Tyr
15 35 40 45
Met Gln His Pro Gly Lys Val Ile Lys Gln Asn Asn Lys Tyr Tyr Phe
50 55 60
Gln Thr Val Leu Asn Asn Ala Ser Phe Trp Lys Glu Tyr Lys Phe Tyr
20 65 70 75 80
Asn Ala Asn Asn Gln Glu Leu Ala Thr Thr Val Val Asn Asp Asn Lys
85 90 95
Lys Ala Asp Thr Arg Thr Ile Asn Val Ala Val Glu Pro Gly Tyr Lys
25 100 105 110
Ser Leu Thr Thr Lys Val His Ile Val Val Pro Gln Ile Asn Tyr Asn
115 120 125
His Arg Tyr Thr Thr His Leu Glu Phe Glu Lys Ala Ile Pro Thr Leu
30 130 135 140
Ala Asp Ala Ala Lys Pro Asn Asn Val Lys Pro Val Gln Pro Lys Pro
145 150 155 160
Ala Gln Pro Lys Thr Pro Thr Glu Gln Thr Lys Pro Val Gln Pro Lys
35 165 170 175
Val Glu Lys Val Lys Pro Thr Val Thr Thr Thr Ser Lys Val Glu Asp
180 185 190
Asn His Ser Thr Lys Val Val Ser Thr Asp Thr Thr Lys Asp Gln Thr
40 195 200 205
Lys Thr Gln Thr Ala His Thr Val Lys Thr Ala Gln Thr Ala Gln Glu
210 215 220
Gln Asn Lys Val Gln Thr Pro Val Lys Asp Val Ala Thr Ala Lys Ser
45 225 230 235 240
Glu Ser Asn Asn Gln Ala Val Ser Asp Asn Lys Ser Gln Gln Thr Asn
245 250 255
Lys Val Thr Lys His Asn Glu Thr Pro Lys Gln Ala Ser Lys Ala Lys
260 265 270
50 Glu Ala Ser Gly Gly Gly Ser Met Ala Met Ile Lys Met Ser Pro Glu
275 280 285
Glu Ile Arg Ala Lys Ser Gln Ser Tyr Gly Gln Gly Ser Asp Gln Ile
55 290 295 300
Arg Gln Ile Leu Ser Asp Leu Thr Arg Ala Gln Gly Glu Ile Ala Ala
305 310 315 320

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Asn Trp Glu Gly Gln Ala Phe Ser Arg Phe Glu Glu Gln Phe Gln Gln
325 330 335

5 Leu Ser Pro Lys Val Glu Lys Phe Ala Gln Leu Leu Glu Glu Ile Lys
340 345 350

Gln Gln Leu Asn Ser Thr Ala Asp Ala Val Gln Glu Gln Asp Gln Gln
355 360 365

10 Leu Ser Asn Asn Phe Gly Leu Gln Ala Ser Gly Gly Gly Ser Met Gly
370 375 380

Gly Tyr Lys Gly Ile Lys Ala Asp Gly Gly Lys Val Asp Gln Ala Lys
385 390 395 400

15 Gln Leu Ala Ala Lys Thr Ala Lys Asp Ile Glu Ala Cys Gln Lys Gln
405 410 415

Thr Gln Gln Leu Ala Glu Tyr Ile Glu Gly Ser Asp Trp Glu Gly Gln
420 425 430

20 Phe Ala Asn Lys Val Lys Asp Val Leu Leu Ile Met Ala Lys Phe Gln
435 440 445

Glu Glu Leu Val Gln Pro Met Ala Asp His Gln Lys Ala Ile Asp Asn
450 455 460

25 Leu Ser Gln Asn Leu Ala Lys Tyr Asp Thr Leu Ser Ile Lys Gln Gly
465 470 475 480

Leu Asp Arg Val Asn Pro
485

30 <210> 213
<211> 256
<212> PRT
<213> Staphylococcus aureus

<400> 213
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35 Leu Val Ile Ser Ile Thr Ala Gly Cys Gly Ile Gly Lys Glu Ala Glu
20 25 30

Val Lys Lys Ser Phe Glu Lys Thr Leu Ser Met Tyr Pro Ile Lys Asn
35 40 45

40 Leu Glu Asp Leu Tyr Asp Lys Glu Gly Tyr Arg Asp Asp Gln Phe Asp
50 55 60

Lys Asn Asp Lys Gly Thr Trp Ile Ile Asn Ser Glu Met Val Ile Gln
65 70 75 80

45 Pro Asn Asn Glu Asp Met Val Ala Lys Gly Met Val Leu Tyr Met Asn
85 90 95

Arg Asn Thr Lys Thr Thr Asn Gly Tyr Tyr Tyr Val Asp Val Thr Lys
100 105 110

50 Asp Glu Asp Glu Gly Lys Pro His Asp Asn Glu Lys Arg Tyr Pro Val
115 120 125

Lys Met Val Asp Asn Lys Ile Ile Pro Thr Lys Glu Ile Lys Asp Glu
130 135 140

55 Lys Leu Lys Lys Glu Ile Glu Asn Phe Lys Phe Phe Val Gln Tyr Gly
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Asp Phe Lys Asn Ile Lys Asn Tyr Lys Asp Gly Asp Ile Ser Tyr Asn

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 <213> Staphylococcus aureus

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Leu Val Ile Ser Ile Thr Ala Gly Cys Gly Ile Gly Lys Glu Ala Glu
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15

Val Lys Lys Ser Phe Glu Lys Thr Leu Ser Met Tyr Pro Ile Lys Asn
 35 40 45

Leu Glu Asp Leu Tyr Asp Lys Glu Gly Tyr Arg Asp Asp Gln Phe Asp
 50 55 60

20

Lys Asn Asp Lys Gly Thr Trp Ile Ile Asn Ser Glu Met Val Ile Gln
 65 70 75 80

Pro Asn Asn Glu Asp Met Val Ala Lys Gly Met Val Leu Tyr Met Asn
 85 90 95

25

Arg Asn Thr Lys Thr Thr Asn Gly Tyr Tyr Tyr Val Asp Val Thr Lys
 100 105 110

Asp Glu Asp Glu Gly Lys Pro His Asp Asn Glu Lys Arg Tyr Pro Val
 115 120 125

30

Lys Met Val Asp Asn Lys Ile Ile Pro Thr Lys Glu Ile Lys Asp Glu
 130 135 140

Lys Leu Lys Lys Glu Ile Glu Asn Phe Lys Phe Phe Val Gln Tyr Gly
 145 150 155 160

Asp Phe Lys Asn Val Lys Asn Tyr Lys Asp Gly Asp Ile Ser Tyr Asn
 165 170 175

35

Pro Glu Val Pro Ser Tyr Ser Ala Lys Tyr Gln Leu Thr Asn Asp Asp
 180 185 190

Tyr Asn Val Lys Gln Leu Arg Lys Arg Tyr Asp Ile Pro Thr Ser Lys
 195 200 205

40

Ala Pro Lys Leu Leu Leu Lys Gly Ser Gly Asn Leu Lys Gly Ser Ser
 210 215 220

Val Gly Tyr Lys Asp Ile Glu Phe Thr Phe Val Glu Lys Lys Glu Glu
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Asn Ile Tyr Phe Ser Asp Ser Leu Asp Tyr Lys Lys Ser Gly Asp Val
 245 250 255

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<210> 216
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 <212> PRT
 <213> Staphylococcus aureus

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Asn Thr Thr Val Lys Thr Gly Asp Leu Val Thr Tyr Asp Lys Glu Asn
 20 25 30

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Gly Met His Lys Lys Val Phe Tyr Ser Phe Ile Asp Asp Lys Asn His
 35 40 45

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Asn Lys Lys Leu Leu Val Ile Arg Thr Lys Gly Thr Ile Ala Gly Gln
 50 55 60
 Tyr Arg Val Tyr Ser Glu Glu Gly Ala Asn Lys Ser Gly Leu Ala Trp
 65 70 75 80
 Pro Ser Ala Phe Lys Val Gln Leu Gln Leu Pro Asp Asn Glu Val Ala
 85 90 95
 Gln Ile Ser Asp Tyr Tyr Pro Arg Asn Ser Ile Asp Thr Pro Ser Gly
 100 105 110
 Ser Val Gln Pro Asp Phe Lys Thr Ile Leu Glu Ser Pro Thr Asp Lys
 115 120 125
 Lys Val Gly Trp Lys Val Ile Phe Asn Asn Met Val Asn Gln Asn Trp
 130 135 140
 Gly Pro Tyr Asp Arg Asp Ser Trp Asn Pro Val Tyr Gly Asn Gln Leu
 145 150 155 160
 Phe Met Lys Thr Arg Asn Gly Ser Met Lys Ala Ala Asp Asn Phe Leu
 165 170 175
 Asp Pro Asn Lys Ala Ser Ser Leu Leu Ser Ser Gly Phe Ser Pro Asp
 180 185 190
 Phe Ala Thr Val Ile Thr Met Asp Arg Lys Ala Ser Lys Gln Gln Thr
 195 200 205
 Asn Ile Asp Val Ile Tyr Glu Arg Val Arg Asp Asp Tyr Gln Leu His
 210 215 220
 Trp Thr Ser Thr Asn Trp Lys Gly Thr Asn Thr Lys Asp Lys Trp Ile
 225 230 235 240
 Asp Arg Ser Ser Glu Arg Tyr Lys Ile Asp Trp Glu Lys Glu Glu Met
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 Thr Asn
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 Gly Asp Asp Thr Gly Lys Ile Gly Gly Leu Ile Gly Ala Asn Val Ser
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 Ile Gly His Thr Leu Lys Tyr Val
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 <212> PRT
 <213> Staphylococcus aureus
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 Gly Met His Lys Lys Val Phe Tyr Ser Phe Ile Asp Asp Lys Asn His

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35 40 45
 Asn Lys Lys Leu Leu Val Ile Arg Thr Lys Gly Thr Ile Ala Gly Gln
 50 55 60
 5 Tyr Arg Val Tyr Ser Glu Glu Gly Ala Asn Lys Ser Gly Leu Ala Trp
 65 70 75 80
 Pro Ser Ala Phe Lys Val Gln Leu Gln Leu Pro Asp Asn Glu Val Ala
 85 90 95
 10 Gln Ile Ser Asp Tyr Tyr Pro Arg Asn Ser Ile Asp Thr
 100 105
 <210> 219
 <211> 145
 <212> PRT
 15 <213> Staphylococcus aureus
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 20 25 30
 Pro Tyr Asp Arg Asp Ser Trp Asn Pro Val Tyr Gly Asn Gln Leu Phe
 35 40 45
 25 Met Lys Thr Arg Asn Gly Ser Met Lys Ala Ala Asp Asn Phe Leu Asp
 50 55 60
 Pro Asn Lys Ala Ser Ser Leu Leu Ser Ser Gly Phe Ser Pro Asp Phe
 65 70 75 80
 30 Ala Thr Val Ile Thr Met Asp Arg Lys Ala Ser Lys Gln Gln Thr Asn
 85 90 95
 Ile Asp Val Ile Tyr Glu Arg Val Arg Asp Asp Tyr Gln Leu His Trp
 100 105 110
 35 Thr Ser Thr Asn Trp Lys Gly Thr Asn Thr Lys Asp Lys Trp Ile Asp
 115 120 125
 Arg Ser Ser Glu Arg Tyr Lys Ile Asp Trp Glu Lys Glu Glu Met Thr
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 35 40 45
 55 Asn Lys Lys Leu Leu Val Ile Arg Thr Lys Gly Thr Ile Ala Gly Gln
 50 55 60
 Tyr Arg Val Tyr Ser Glu Glu Gly Ala Asn Lys Ser Gly Leu Ala Trp
 65 70 75 80

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Pro Ser Ala Phe Lys Val Gln Leu Gln Leu Pro Asp Asn Glu Val Ala
 85 90 95
 Gln Ile Ser Asp Tyr Tyr Pro Arg Asn Ser Ile Asp Thr Lys Glu Tyr
 100 105 110
 Met Ser Thr Leu Thr Tyr Gly Phe Asn Gly Asn Val Thr Gly Asp Asp
 115 120 125
 Thr Gly Lys Ile Gly Gly Leu Ile Gly Ala Asn Val Ser Ile Gly His
 130 135 140
 Thr Leu Lys Tyr Val Gln Pro Asp Phe Lys Thr Ile Leu Glu Ser Pro
 145 150 155 160
 Thr Asp Lys Lys Val Gly Trp Lys Val Ile Phe Asn Asn Met Val Asn
 165 170 175
 Gln Asn Trp Gly Pro Tyr Asp Arg Asp Ser Trp Asn Pro Val Tyr Gly
 180 185 190
 Asn Gln Leu Phe Met Lys Thr Arg Asn Gly Ser Met Lys Ala Ala Asp
 195 200 205
 Asn Phe Leu Asp Pro Asn Lys Ala Ser Ser Leu Leu Ser Ser Gly Phe
 210 215 220
 Ser Pro Asp Phe Ala Thr Val Ile Thr Met Asp Arg Lys Ala Ser Lys
 225 230 235 240
 Gln Gln Thr Asn Ile Asp Val Ile Tyr Glu Arg Val Arg Asp Asp Tyr
 245 250 255
 Gln Leu His Trp Thr Ser Thr Asn Trp Lys Gly Thr Asn Thr Lys Asp
 260 265 270
 Lys Trp Ile Asp Arg Ser Ser Glu Arg Tyr Lys Ile Asp Trp Glu Lys
 275 280 285
 Glu Glu Met Thr Asn Ala Ser Gly Gly Gly Ser Met Ala Met Ile Lys
 290 295 300
 Met Ser Pro Glu Glu Ile Arg Ala Lys Ser Gln Ser Tyr Gly Gln Gly
 305 310 315 320
 Ser Asp Gln Ile Arg Gln Ile Leu Ser Asp Leu Thr Arg Ala Gln Gly
 325 330 335
 Glu Ile Ala Ala Asn Trp Glu Gly Gln Ala Phe Ser Arg Phe Glu Glu
 340 345 350
 Gln Phe Gln Gln Leu Ser Pro Lys Val Glu Lys Phe Ala Gln Leu Leu
 355 360 365
 Glu Glu Ile Lys Gln Gln Leu Asn Ser Thr Ala Asp Ala Val Gln Glu
 370 375 380
 Gln Asp Gln Gln Leu Ser Asn Asn Phe Gly Leu Gln Ala Ser Gly Gly
 385 390 395 400
 Gly Ser Met Gly Gly Tyr Lys Gly Ile Lys Ala Asp Gly Gly Lys Val
 405 410 415
 Asp Gln Ala Lys Gln Leu Ala Ala Lys Thr Ala Lys Asp Ile Glu Ala
 420 425 430
 Cys Gln Lys Gln Thr Gln Gln Leu Ala Glu Tyr Ile Glu Gly Ser Asp
 435 440 445

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Trp Glu Gly Gln Phe Ala Asn Lys Val Lys Asp Val Leu Leu Ile Met
 450 455 460
 5 Ala Lys Phe Gln Glu Glu Leu Val Gln Pro Met Ala Asp His Gln Lys
 465 470 475
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 485 490 495
 10 Ile Lys Gln Gly Leu Asp Arg Val Asn Pro
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 Gln Val Gln Lys Asp Gly Ser Ser Glu Lys Ser His Met Asp Asp Tyr
 35 40 45
 Met Gln His Pro Gly Lys Val Ile Lys Gln Asn Asn Lys Tyr Tyr Phe
 50 55 60
 25 Gln Thr Val Leu Asn Asn Ala Ser Phe Trp Lys Glu Tyr Lys Phe Tyr
 65 70 75 80
 Asn Ala Asn Asn Gln Glu Leu Ala Thr Thr Val Val Asn Asp Asn Lys
 85 90 95
 30 Lys Ala Asp Thr Arg Thr Ile Asn Val Ala Val Glu Pro Gly Tyr Lys
 100 105 110
 Ser Leu Thr Thr Lys Val His Ile Val Val Pro Gln Ile Asn Tyr Asn
 115 120 125
 35 His Arg Tyr Thr Thr His Leu Glu Phe Glu Lys Ala Ile Pro Thr Leu
 130 135 140
 Ala Ala Ser Gly Gly Gly Ser Cys Gly Asn Gln Gly Glu Lys Asn Asn
 145 150 155 160
 40 Lys Ala Glu Thr Lys Ser Tyr Lys Met Asp Asp Gly Lys Thr Val Asp
 165 170 175
 Ile Pro Lys Asp Pro Lys Arg Ile Ala Val Val Ala Pro Thr Tyr Ala
 180 185 190
 45 Gly Gly Leu Lys Lys Leu Gly Ala Asn Ile Val Ala Val Asn Gln Gln
 195 200 205
 Val Asp Gln Ser Lys Val Leu Lys Asp Lys Phe Lys Gly Val Thr Lys
 210 215 220
 50 Ile Gly Asp Gly Asp Val Glu Lys Val Ala Lys Glu Lys Pro Asp Leu
 225 230 235 240
 Ile Ile Val Tyr Ser Thr Asp Lys Asp Ile Lys Lys Tyr Gln Lys Val
 245 250 255
 55 Ala Pro Thr Val Val Val Asp Tyr Asn Lys His Lys Tyr Leu Glu Gln
 260 265 270
 Gln Glu Met Leu Gly Lys Ile Val Gly Lys Glu Asp Lys Val Lys Ala

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		275					280					285				
	Trp	Lys	Lys	Asp	Trp	Glu	Glu	Thr	Thr	Ala	Lys	Asp	Gly	Lys	Glu	Ile
		290					295					300				
5	Lys	Lys	Ala	Ile	Gly	Gln	Asp	Ala	Thr	Val	Ser	Leu	Phe	Asp	Glu	Phe
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	Asp	Lys	Lys	Leu	Tyr	Thr	Tyr	Gly	Asp	Asn	Trp	Gly	Arg	Gly	Gly	Glu
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 515 520 525
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 Ala Glu Tyr Ile Glu Gly Ser Asp Trp Glu Gly Gln Phe Ala Asn Lys
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 25 Val Lys Asp Val Leu Leu Ile Met Ala Lys Phe Gln Glu Glu Leu Val
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 Gln Pro Met Ala Asp His Gln Lys Ala Ile Asp Asn Leu Ser Gln Asn
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 65 70 75 80
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 Thr Leu Lys Tyr Val Gln Pro Asp Phe Lys Thr Ile Leu Glu Ser Pro
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 Thr Asp Lys Lys Val Gly Trp Lys Val Ile Phe Asn Asn Met Val Asn
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 180 185 190
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 195 200 205
 Asn Phe Leu Asp Pro Asn Lys Ala Ser Ser Leu Leu Ser Ser Gly Phe
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 Ser Pro Asp Phe Ala Thr Val Ile Thr Met Asp Arg Lys Ala Ser Lys
 225 230 235 240
 Gln Gln Thr Asn Ile Asp Val Ile Tyr Glu Arg Val Arg Asp Asp Tyr
 245 250 255
 Gln Leu His Trp Thr Ser Thr Asn Trp Lys Gly Thr Asn Thr Lys Asp
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 Lys Glu Tyr Met Ser Thr Leu Thr Tyr Gly Phe Asn Gly Asn Val Thr
 115 120 125
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 130 135 140
 Ile Gly His Thr Leu Lys Tyr Val Gln Pro Asp Phe Lys Thr Ile Leu
 145 150 155 160
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 Glu Ser Pro Thr Asp Lys Lys Val Gly Trp Lys Val Ile Phe Asn Asn
 165 170 175
 Met Val Asn Gln Asn Trp Gly Pro Tyr Asp Arg Asp Ser Trp Asn Pro
 180 185 190
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 Val Tyr Gly Asn Gln Leu Phe Met Lys Thr Arg Asn Gly Ser Met Lys
 195 200 205
 Ala Ala Asp Asn Phe Leu Asp Pro Asn Lys Ala Ser Ser Leu Leu Ser
 210 215 220
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Pro Lys Asn Pro Lys Arg Ile Ala Val Val Ala Pro Ser Tyr Ala Gly
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Gly Ile Lys Lys Leu Gly Gly Asn Val Val Ala Val Ser Asn Gln Val
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Asp Gln Ser Ser Ile Leu Lys Asp Lys Phe Lys Gly Val Thr Lys Val
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Gly Asp Asp Asp Val Glu Lys Val Ala Lys Gln Lys Pro Asp Leu Ile
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Ile Val Leu Asp Gln Asn Lys Asn Ile Lys Lys Tyr Lys Lys Ile Ala
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Ala Thr Val Pro Phe Asn Tyr Gln Lys His Lys Tyr Leu Glu Gln Gln
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Glu Lys Gln Trp Lys Asp Gln Thr Ala Lys Asp Gly Lys Glu Ile Lys
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195 200 205
Leu Tyr Gln Ala Phe Gly Leu Lys Met Ser Lys Gly Gln His Asp Leu
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Thr Lys Lys Ala Gly Trp Ala Glu Val Asn Gln Glu Gln Ile Glu Lys
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Val Ala Gly Asp Tyr Ile Val Ser Thr Ser Ala Gly Lys Ser Thr Pro
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 Asn Gln Leu Phe Met Lys Thr Arg Asn Gly Ser Met Lys Ala Ala Asp
 195 200 205
 10 Asn Phe Leu Asp Pro Asn Lys Ala Ser Ser Leu Leu Ser Ser Gly Phe
 210 215 220
 Ser Pro Asp Phe Ala Thr Val Ile Thr Met Asp Arg Lys Ala Ser Lys
 225 230 235 240
 15 Gln Gln Thr Asn Ile Asp Val Ile Tyr Glu Arg Val Arg Asp Asp Tyr
 245 250 255
 Gln Leu His Trp Thr Ser Thr Asn Trp Lys Gly Thr Asn Thr Lys Asp
 260 265 270
 20 Lys Trp Ile Asp Arg Ser Ser Glu Arg Tyr Lys Ile Asp Trp Glu Lys
 275 280 285
 Glu Glu Met Thr Asn Ala Pro Thr Ala Arg Gly Ala Met Ile Lys Met
 290 295 300
 25 Ser Pro Glu Glu Ile Arg Ala Lys Ser Gln Ser Tyr Gly Gln Gly Ser
 305 310 315 320
 Asp Gln Ile Arg Gln Ile Leu Ser Asp Leu Thr Arg Ala Gln Gly Glu
 325 330 335
 30 Ile Ala Ala Asn Trp Glu Gly Gln Ala Phe Ser Arg Phe Glu Glu Gln
 340 345 350
 Phe Gln Gln Leu Ser Pro Lys Val Glu Lys Phe Ala Gln Leu Leu Glu
 355 360 365
 35 Glu Ile Lys Gln Gln Leu Asn Ser Thr Ala Asp Ala Val Gln Glu Gln
 370 375 380
 Asp Gln Gln Leu Ser Asn Asn Phe Gly Leu Gln Ala Ser Gly Gly Gly
 385 390 395 400
 40 Ser Gly Gly Tyr Lys Gly Ile Lys Ala Asp Gly Gly Lys Val Asp Gln
 405 410 415
 Ala Lys Gln Leu Ala Ala Lys Thr Ala Lys Asp Ile Glu Ala Cys Gln
 420 425 430
 45 Lys Gln Thr Gln Gln Leu Ala Glu Tyr Ile Glu Gly Ser Asp Trp Glu
 435 440 445
 Gly Gln Phe Ala Asn Lys Val Lys Asp Val Leu Leu Ile Met Ala Lys
 450 455 460
 50 Phe Gln Glu Glu Leu Val Gln Pro Met Ala Asp His Gln Lys Ala Ile
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 <213> Staphylococcus aureus

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 5 Lys Met Asp Asp Gly Lys Thr Val Asp Ile Pro Lys Asp Pro Lys Arg
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 15 Lys Val Ala Lys Glu Lys Pro Asp Leu Ile Ile Val Tyr Ser Thr Asp
 20 Lys Asp Ile Lys Lys Tyr Gln Lys Val Ala Pro Thr Val Val Val Asp
 25 Tyr Asn Lys His Lys Tyr Leu Glu Gln Gln Glu Met Leu Gly Lys Ile
 30 Val Gly Lys Glu Asp Lys Val Lys Ala Trp Lys Lys Asp Trp Glu Glu
 35 Thr Thr Ala Lys Asp Gly Lys Glu Ile Lys Lys Ala Ile Gly Gln Asp
 40 Ala Thr Val Ser Leu Phe Asp Glu Phe Asp Lys Lys Leu Tyr Thr Tyr
 45 Gly Asp Asn Trp Gly Arg Gly Gly Glu Val Leu Tyr Gln Ala Phe Gly
 50 Leu Lys Met Gln Pro Glu Gln Gln Lys Leu Thr Ala Lys Ala Gly Trp
 55 Ala Glu Val Lys Gln Glu Glu Ile Glu Lys Tyr Ala Gly Asp Tyr Ile
 60 Val Ser Thr Ser Glu Gly Lys Pro Thr Pro Gly Tyr Glu Ser Thr Asn
 65 Met Trp Lys Asn Leu Lys Ala Thr Lys Glu Gly His Ile Val Lys Val
 70 Asp Ala Gly Thr Tyr Trp Tyr Asn Asp Pro Tyr Thr Leu Asp Phe Met
 75 Arg Lys Asp Leu Lys Glu Lys Leu Ile Lys Ala Ala Lys Ala Pro Thr
 80 Ala Arg Gly Ala Met Ile Lys Met Ser Pro Glu Glu Ile Arg Ala Lys
 85 Ser Gln Ser Tyr Gly Gln Gly Ser Asp Gln Ile Arg Gln Ile Leu Ser
 90 Asp Leu Thr Arg Ala Gln Gly Glu Ile Ala Ala Asn Trp Glu Gly Gln
 95 Ala Phe Ser Arg Phe Glu Glu Gln Phe Gln Gln Leu Ser Pro Lys Val
 100 Glu Lys Phe Ala Gln Leu Leu Glu Glu Ile Lys Gln Gln Leu Asn Ser

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Thr Ala Asp Ala Val Gln Glu Gln Asp Gln Gln Leu Ser Asn Asn Phe
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 405 410 415
 10 Ala Lys Asp Ile Glu Ala Cys Gln Lys Gln Thr Gln Gln Leu Ala Glu
 420 425 430
 Tyr Ile Glu Gly Ser Asp Trp Glu Gly Gln Phe Ala Asn Lys Val Lys
 435 440 445
 15 Asp Val Leu Leu Ile Met Ala Lys Phe Gln Glu Glu Leu Val Gln Pro
 450 455 460
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 35 40 45
 Arg Phe Glu Glu Gln Phe Gln Gln Leu Ser Pro Lys Val Glu Lys Phe
 50 55 60
 55 Ala Gln Leu Leu Glu Glu Ile Lys Gln Gln Leu Asn Ser Thr Ala Asp
 65 70 75 80

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Ala Val Gln Glu Gln Asp Gln Gln Leu Ser Asn Asn Phe Gly Leu Gln
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Ala Ser Gly Gly Gly Ser Gly Gly Tyr Lys Gly Ile Lys Ala Asp Gly
5 100 105 110

Gly Lys Val Asp Gln Ala Lys Gln Leu Ala Ala Lys Thr Ala Lys Asp
115 120 125

Ile Glu Ala Cys Gln Lys Gln Thr Gln Gln Leu Ala Glu Tyr Ile Glu
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Gly Ser Asp Trp Glu Gly Gln Phe Ala Asn Lys Val Lys Asp Val Leu
145 150 155 160

Leu Ile Met Ala Lys Phe Gln Glu Glu Leu Val Gln Pro Met Ala Asp
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20 25 30

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35 40 45

Asn Lys Lys Leu Leu Val Ile Arg Thr Lys Gly Thr Ile Ala Gly Gln
50 55 60

Tyr Arg Val Tyr Ser Glu Glu Gly Ala Asn Lys Ser Gly Leu Ala Trp
65 70 75 80

Pro Ser Ala Phe Lys Val Gln Leu Gln Leu Pro Asp Asn Glu Val Ala
85 90 95

40
Gln Ile Ser Asp Leu Tyr Pro Arg Asn Ser Ile Asp Thr Lys Glu Tyr
100 105 110

Met Ser Thr Leu Thr Tyr Gly Phe Asn Gly Asn Val Thr Gly Asp Asp
115 120 125

45
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Thr Leu Lys Tyr Val Gln Pro Asp Phe Lys Thr Ile Leu Glu Ser Pro
145 150 155 160

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Thr Asp Lys Lys Val Gly Trp Lys Val Ile Phe Asn Asn Met Val Asn
165 170 175

Gln Asn Trp Gly Pro Tyr Asp Arg Asp Ser Trp Asn Pro Val Tyr Gly
180 185 190

55
Asn Gln Leu Phe Met Lys Thr Arg Asn Gly Ser Met Lys Ala Ala Asp
195 200 205

Asn Phe Leu Asp Pro Asn Lys Ala Ser Ser Leu Leu Ser Ser Gly Phe

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	Glu	Glu 290	Met	Thr	Asn												
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	<211>	293															
	<212>	PRT															
	<213>	Staphylococcus aureus															
	<400>	243															
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25	Gly	Met	His 35	Lys	Lys	Val	Phe	Tyr 40	Ser	Phe	Ile	Asp 45	Lys	Asn	His		
	Asn	Lys 50	Lys	Leu	Leu	Val	Ile 55	Arg	Thr	Lys	Gly	Thr 60	Ile	Ala	Gly	Gln	
30	Tyr 65	Arg	Val	Tyr	Ser	Glu 70	Glu	Gly	Ala	Asn	Lys 75	Ser	Gly	Leu	Ala	Trp 80	
	Pro	Ser	Ala	Phe	Lys 85	Val	Gln	Leu	Gln	Leu 90	Pro	Asp	Asn	Glu	Val 95	Ala	
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	Met	Ser	Thr 115	Leu	Thr	Tyr	Gly	Phe 120	Asn	Gly	Asn	Val	Thr 125	Gly	Asp	Asp	
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	Thr 145	Leu	Lys	Tyr	Val	Gln 150	Pro	Leu	Phe	Lys	Thr 155	Ile	Leu	Glu	Ser	Pro 160	
45	Thr	Asp	Lys	Lys	Val 165	Gly	Trp	Lys	Val	Ile 170	Phe	Asn	Asn	Met	Val 175	Asn	
	Gln	Asn	Trp	Gly 180	Pro	Tyr	Asp	Arg	Asp 185	Ser	Trp	Asn	Pro	Val 190	Tyr	Gly	
50	Asn	Gln	Leu 195	Phe	Met	Lys	Thr	Arg 200	Asn	Gly	Ser	Met	Lys 205	Ala	Ala	Asp	
	Asn	Phe 210	Leu	Asp	Pro	Asn	Lys 215	Ala	Ser	Ser	Leu	Leu 220	Ser	Ser	Gly	Phe	
55	Ser 225	Pro	Asp	Phe	Ala	Thr 230	Val	Ile	Thr	Met	Asp 235	Arg	Lys	Ala	Ser	Lys 240	
	Gln	Gln	Thr	Asn	Ile 245	Asp	Val	Ile	Tyr	Glu 250	Arg	Val	Arg	Asp	Asp 255	Tyr	

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Gln Leu His Trp Thr Ser Thr Asn Trp Lys Gly Thr Asn Thr Lys Asp
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 Glu Glu Met Thr Asn
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 <211> 293
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 <213> Staphylococcus aureus
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 20 35 40 45
 Asn Lys Lys Leu Leu Val Ile Arg Thr Lys Gly Thr Ile Ala Gly Gln
 50 55 60
 Tyr Arg Val Tyr Ser Glu Glu Gly Ala Asn Lys Ser Gly Leu Ala Trp
 25 65 70 75 80
 Pro Ser Ala Phe Lys Val Gln Leu Gln Leu Pro Asp Asn Glu Val Ala
 85 90 95
 Gln Ile Ser Asp Leu Tyr Pro Arg Asn Ser Ile Asp Thr Lys Glu Tyr
 30 100 105 110
 Met Ser Thr Leu Thr Tyr Gly Phe Asn Gly Asn Val Thr Gly Asp Asp
 115 120 125
 Thr Gly Lys Ile Gly Gly Leu Ile Gly Ala Asn Val Ser Ile Gly His
 35 130 135 140
 Thr Leu Lys Tyr Val Gln Pro Asp Phe Lys Thr Ile Leu Glu Ser Pro
 145 150 155 160
 Thr Asp Lys Lys Val Gly Trp Lys Val Ile Phe Asn Asn Met Val Asn
 40 165 170 175
 Gln Asn Trp Gly Pro Tyr Asp Arg Asp Ser Trp Asn Pro Val Tyr Gly
 180 185 190
 Asn Gln Leu Phe Met Lys Thr Arg Asn Gly Ser Met Lys Ala Ala Asp
 45 195 200 205
 Asn Phe Leu Asp Pro Asn Lys Ala Ser Ser Leu Leu Ser Ser Gly Phe
 210 215 220
 Ser Pro Asp Phe Ala Thr Val Ile Thr Met Asp Arg Lys Ala Ser Lys
 50 225 230 235 240
 Gln Gln Thr Asn Ile Asp Val Ile Tyr Glu Arg Val Arg Asp Asp Tyr
 245 250 255
 Gln Leu His Trp Thr Ser Thr Asn Trp Lys Gly Thr Asn Thr Lys Asp
 260 265 270
 Lys Trp Ile Asp Arg Ser Ser Glu Arg Tyr Lys Ile Asp Trp Glu Lys
 55 275 280 285

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Glu Glu Met Thr Asn
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5
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20 30

Gly Met Leu Lys Lys Val Phe Tyr Ser Phe Ile Asp Asp Lys Asn His
35 40 45

15
Asn Lys Lys Leu Leu Val Ile Arg Thr Lys Gly Thr Ile Ala Gly Gln
50 55 60

Tyr Arg Val Tyr Ser Glu Glu Gly Ala Asn Lys Ser Gly Leu Ala Trp
65 70 75 80

20
Pro Ser Ala Phe Lys Val Gln Leu Gln Leu Pro Asp Asn Glu Val Ala
85 90 95

Gln Ile Ser Asp Tyr Tyr Pro Arg Asn Ser Ile Asp Thr Lys Glu Tyr
100 105 110

25
Met Ser Thr Leu Thr Tyr Gly Phe Asn Gly Asn Val Thr Gly Asp Asp
115 120 125

Thr Gly Lys Ile Gly Gly Leu Ile Gly Ala Asn Val Ser Ile Gly His
130 135 140

30
Thr Leu Lys Tyr Val Gln Pro Leu Phe Lys Thr Ile Leu Glu Ser Pro
145 150 155 160

Thr Asp Lys Lys Val Gly Trp Lys Val Ile Phe Asn Asn Met Val Asn
165 170 175

35
Gln Asn Trp Gly Pro Tyr Asp Arg Asp Ser Trp Asn Pro Val Tyr Gly
180 185 190

Asn Gln Leu Phe Met Lys Thr Arg Asn Gly Ser Met Lys Ala Ala Asp
195 200 205

40
Asn Phe Leu Asp Pro Asn Lys Ala Ser Ser Leu Leu Ser Ser Gly Phe
210 215 220

Ser Pro Asp Phe Ala Thr Val Ile Thr Met Asp Arg Lys Ala Ser Lys
225 230 235 240

45
Gln Gln Thr Asn Ile Asp Val Ile Tyr Glu Arg Val Arg Asp Asp Tyr
245 250 255

Gln Leu His Trp Thr Ser Thr Asn Trp Lys Gly Thr Asn Thr Lys Asp
260 265 270

50
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275 280 285

Glu Glu Met Thr Asn
290

55
<210> 246
<211> 285
<212> PRT
<213> Staphylococcus aureus

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 Ile Ala Val Val Ala Pro Thr Tyr Ala Gly Gly Leu Lys Lys Leu Gly
 35 40 45
 10 Ala Asn Ile Val Ala Val Asn Gln Gln Val Asp Gln Ser Lys Val Leu
 50 55 60
 Lys Asp Lys Phe Lys Gly Val Thr Lys Ile Gly Asp Gly Asp Val Glu
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 15 Lys Val Ala Lys Glu Lys Pro Asp Leu Ile Ile Val Tyr Ser Thr Asp
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 Lys Asp Ile Lys Lys Tyr Gln Lys Val Ala Pro Thr Val Val Val Asp
 100 105 110
 20 Tyr Asn Lys His Lys Tyr Leu Glu Gln Gln Glu Met Leu Gly Lys Ile
 115 120 125
 Val Gly Lys Glu Asp Lys Val Lys Ala Trp Lys Lys Asp Trp Glu Glu
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 25 Thr Thr Ala Lys Asp Gly Lys Glu Ile Lys Lys Ala Ile Gly Gln Asp
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 Ala Thr Val Ser Leu Phe Asp Glu Phe Asp Lys Lys Leu Tyr Thr Tyr
 165 170 175
 30 Gly Asp Asn Trp Gly Arg Gly Gly Glu Val Leu Tyr Gln Ala Phe Gly
 180 185 190
 Leu Lys Met Gln Pro Glu Gln Gln Lys Leu Thr Ala Lys Ala Gly Trp
 195 200 205
 35 Ala Glu Val Lys Gln Glu Glu Ile Glu Lys Tyr Ala Gly Asp Tyr Ile
 210 215 220
 Val Ser Thr Ser Glu Gly Lys Pro Thr Pro Gly Tyr Glu Ser Thr Asn
 225 230 235 240
 40 Met Trp Lys Asn Leu Lys Ala Thr Lys Glu Gly His Ile Val Lys Val
 245 250 255
 Asp Ala Gly Thr Tyr Trp Tyr Asn Asp Pro Tyr Thr Leu Asp Phe Met
 260 265 270
 45 Arg Lys Asp Leu Lys Glu Lys Leu Ile Lys Ala Ala Lys
 275 280 285
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 <211> 233
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 <213> Staphylococcus aureus
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 55 Glu Gly Tyr Arg Asp Asp Gln Phe Asp Lys Asn Asp Lys Gly Thr Trp
 35 40 45

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Ile Ile Asn Ser Glu Met Val Ile Gln Pro Asn Asn Glu Asp Met Val
50 55 60

Ala Lys Gly Met Val Leu Tyr Met Asn Arg Asn Thr Lys Thr Thr Asn
65 70 75 80

Gly Tyr Tyr Tyr Val Asp Val Thr Lys Asp Glu Asp Glu Gly Lys Pro
85 90 95

His Asp Asn Glu Lys Arg Tyr Pro Val Lys Met Val Asp Asn Lys Ile
100 105 110

Ile Pro Thr Lys Glu Ile Lys Asp Glu Lys Ile Lys Lys Glu Ile Glu
115 120 125

Asn Phe Lys Phe Phe Val Gln Tyr Gly Asp Phe Lys Asn Leu Lys Asn
130 135 140

Tyr Lys Asp Gly Asp Ile Ser Tyr Asn Pro Glu Val Pro Ser Tyr Ser
145 150 160

Ala Lys Tyr Gln Leu Thr Asn Asp Asp Tyr Asn Val Lys Gln Leu Arg
165 170 175

Lys Arg Tyr Asp Ile Pro Thr Ser Lys Ala Pro Lys Leu Leu Lys
180 185 190

Gly Ser Gly Asn Leu Lys Gly Ser Ser Val Gly Tyr Lys Asp Ile Glu
195 200 205

Phe Thr Phe Val Glu Lys Lys Glu Glu Asn Ile Tyr Phe Ser Asp Ser
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Leu Asp Tyr Lys Lys Ser Gly Asp Val
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<212> PRT
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20 25 30

Pro Lys Arg Ile Ala Val Val Ala Pro Thr Tyr Ala Gly Gly Leu Lys
35 40 45

Lys Leu Gly Ala Asn Ile Val Ala Val Asn Gln Gln Val Asp Gln Ser
50 55 60

Lys Val Leu Lys Asp Lys Phe Lys Gly Val Thr Lys Ile Gly Asp Gly
65 70 75 80

Asp Val Glu Lys Val Ala Lys Glu Lys Pro Asp Leu Ile Ile Val Tyr
85 90 95

Ser Thr Asp Lys Asp Ile Lys Lys Tyr Gln Lys Val Ala Pro Thr Val
100 105 110

Val Val Asp Tyr Asn Lys His Lys Tyr Leu Glu Gln Gln Glu Met Leu
115 120 125

Gly Lys Ile Val Gly Lys Glu Asp Lys Val Lys Ala Trp Lys Lys Asp
130 135 140

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Trp Glu Glu Thr Thr Ala Lys Asp Gly Lys Glu Ile Lys Lys Ala Ile
 145 150 155 160
 5 Gly Gln Asp Ala Thr Val Ser Leu Phe Asp Glu Phe Asp Lys Lys Leu
 165 170 175
 Tyr Thr Tyr Gly Asp Asn Trp Gly Arg Gly Gly Glu Val Leu Tyr Gln
 180 185 190
 10 Ala Phe Gly Leu Lys Met Gln Pro Glu Gln Gln Lys Leu Thr Ala Lys
 195 200 205
 Ala Gly Trp Ala Glu Val Lys Gln Glu Glu Ile Glu Lys Tyr Ala Gly
 210 215 220
 15 Asp Tyr Ile Val Ser Thr Ser Glu Gly Lys Pro Thr Pro Gly Tyr Glu
 225 230 235
 Ser Thr Asn Met Trp Lys Asn Leu Lys Ala Thr Lys Glu Gly His Ile
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 Lys Glu Gly Tyr Arg Asp Asp Gln Phe Asp Lys Asn Asp Lys Gly Thr
 35 35 40 45
 Trp Ile Ile Asn Ser Glu Met Val Ile Gln Pro Asn Asn Glu Asp Met
 50 55 60
 Val Ala Lys Gly Met Val Leu Tyr Met Asn Arg Asn Thr Lys Thr Thr
 65 70 75 80
 40 Asn Gly Tyr Tyr Tyr Val Asp Val Thr Lys Asp Glu Asp Glu Gly Lys
 85 90 95
 Pro His Asp Asn Glu Lys Arg Tyr Pro Val Lys Met Val Asp Asn Lys
 100 105 110
 45 Ile Ile Pro Thr Lys Glu Ile Lys Asp Glu Lys Ile Lys Lys Glu Ile
 115 120 125
 Glu Asn Phe Lys Phe Phe Val Gln Tyr Gly Asp Phe Lys Asn Leu Lys
 130 135 140
 50 Asn Tyr Lys Asp Gly Asp Ile Ser Tyr Asn Pro Glu Val Pro Ser Tyr
 145 150 155 160
 Ser Ala Lys Tyr Gln Leu Thr Asn Asp Asp Tyr Asn Val Lys Gln Leu
 165 170 175
 55 Arg Lys Arg Tyr Asp Ile Pro Thr Ser Lys Ala Pro Lys Leu Leu Leu
 180 185 190
 Lys Gly Ser Gly Asn Leu Lys Gly Ser Ser Val Gly Tyr Lys Asp Ile

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ID NO: 42, wherein 'n' is 7 or more.

4. The composition of claim 3 for use in a method for raising an immune response in a mammal.

5 5. The composition of any preceding claim, wherein the sta006 antigen comprises an amino acid sequence having 60%, 70%, 80%, 90% or more identity to SEQ ID NO: 42.

10 6. The composition of any preceding claim, wherein the sta006 antigen comprises an amino acid sequence having 90% or more identity to SEQ ID NO: 246.

7. The composition of any preceding claim, wherein the sta006 antigen comprises an amino acid having 99% or more identity to SEQ ID NO: 248.

15 8. The composition of any of claims 2-7, wherein the adjuvant is aluminium hydroxide.

9. The composition of any preceding claim, wherein the composition includes a histidine buffer or a phosphate buffer.

20 10. The composition of any preceding claim, further comprising one or more conjugates of a *S.aureus* exopolysaccharide and a carrier protein.

11. The composition of any of claims 1-9, further comprising one or more conjugates of a *S.aureus* capsular polysaccharide and a carrier protein.

25 12. The composition of any preceding claim, in lyophilized form.

13. The composition of any of claims 1-11, in aqueous form.

30 14. A method for preparing the composition of claim 13, by reconstituting the composition of claim 12 with aqueous material.

35 15. A pharmaceutical composition comprising the composition of any preceding claim, and a pharmaceutical carrier and/or an excipient.

40

45

50

55

FIGURE 1

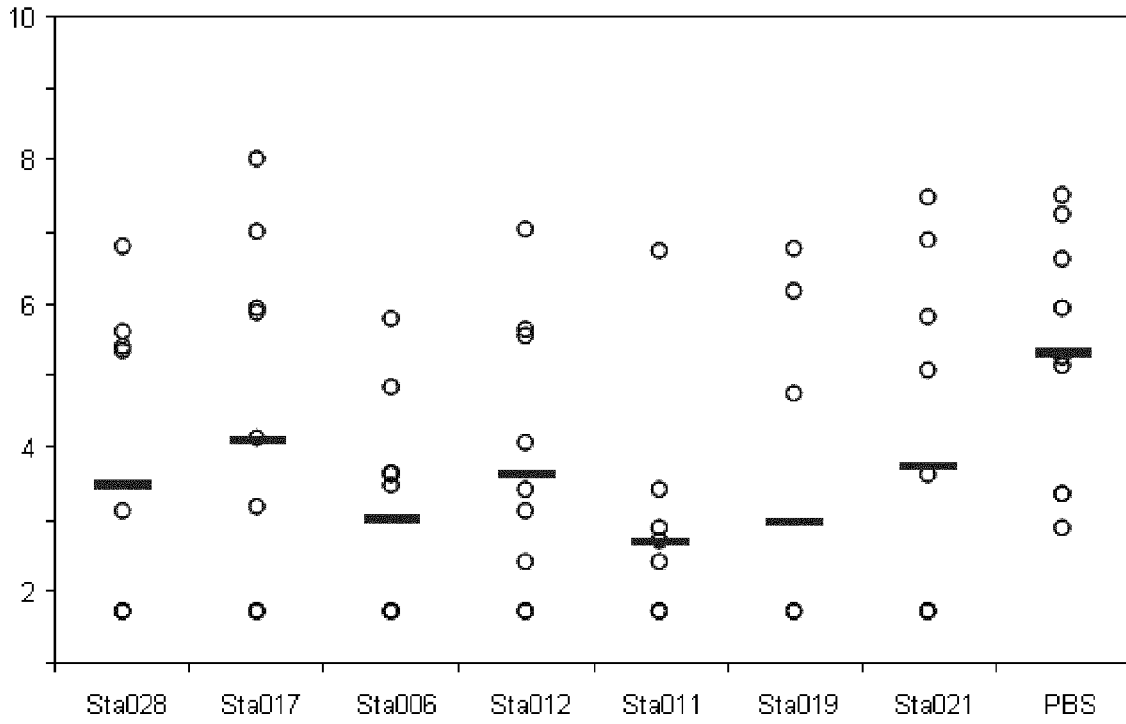


FIGURE 2

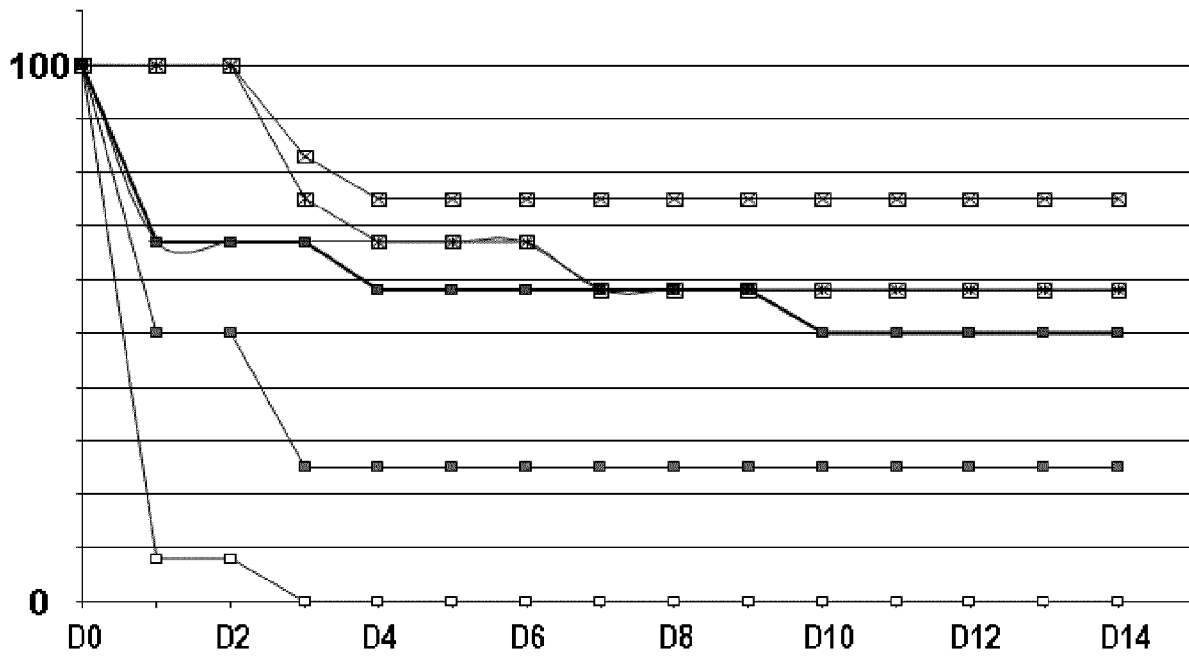


FIGURE 3

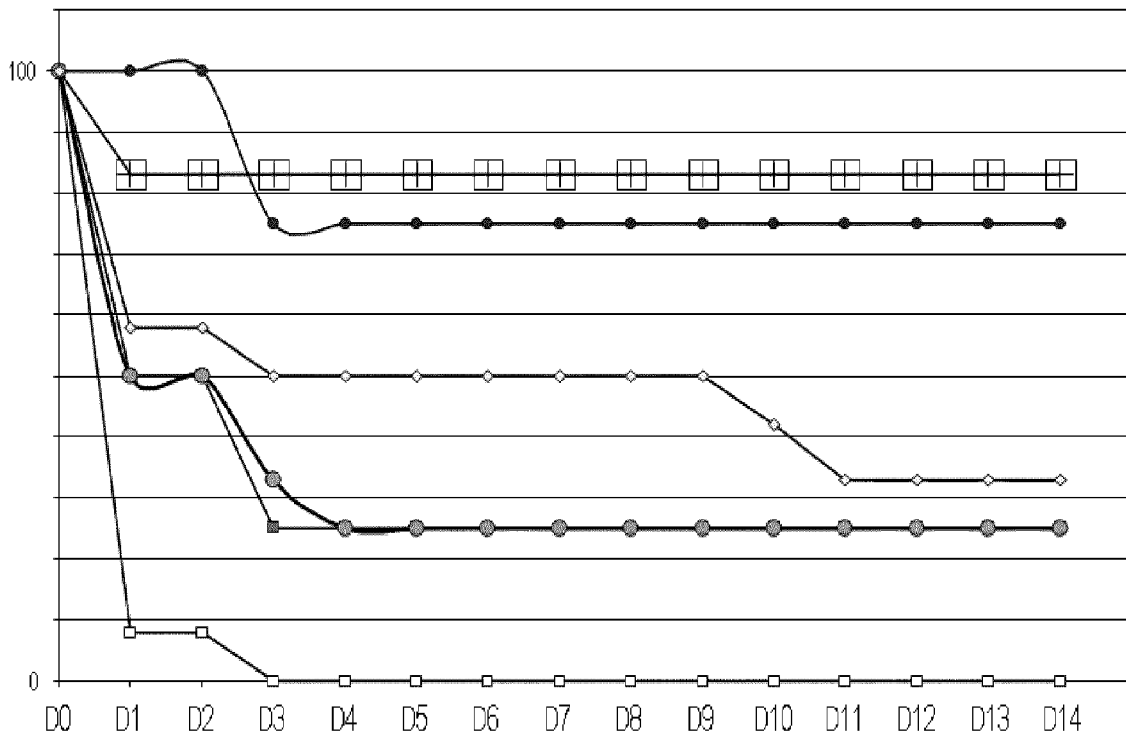


FIGURE 4

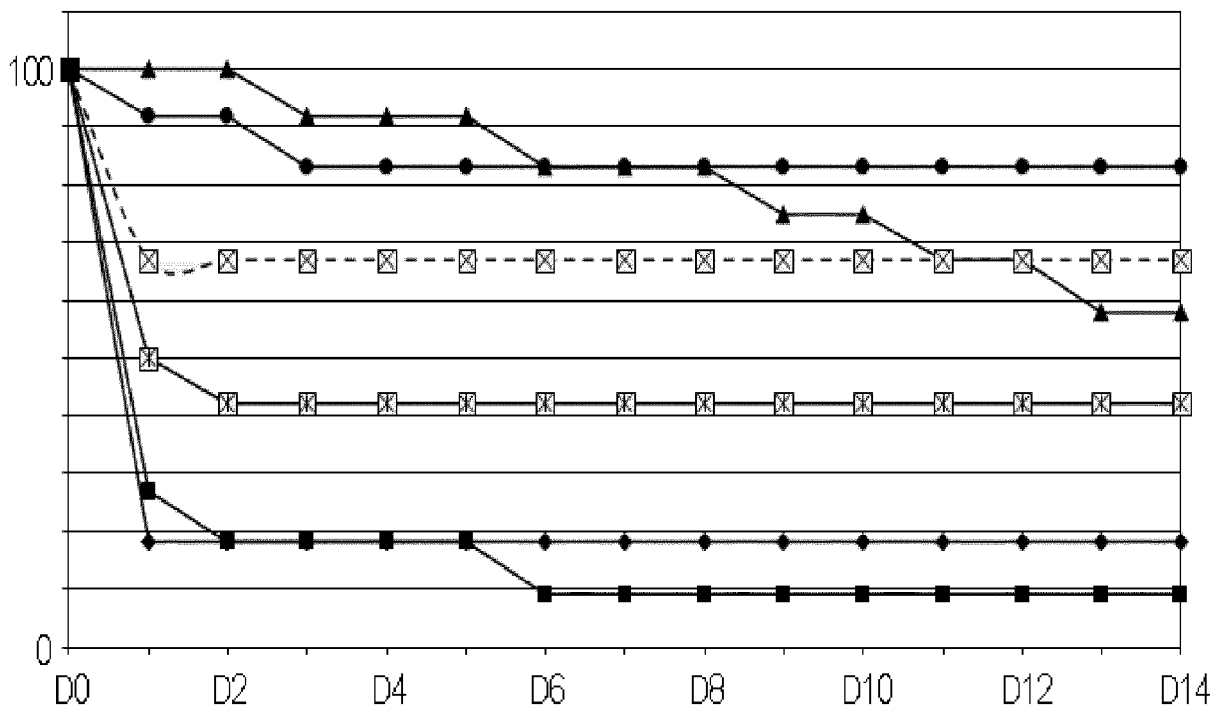


FIGURE 5

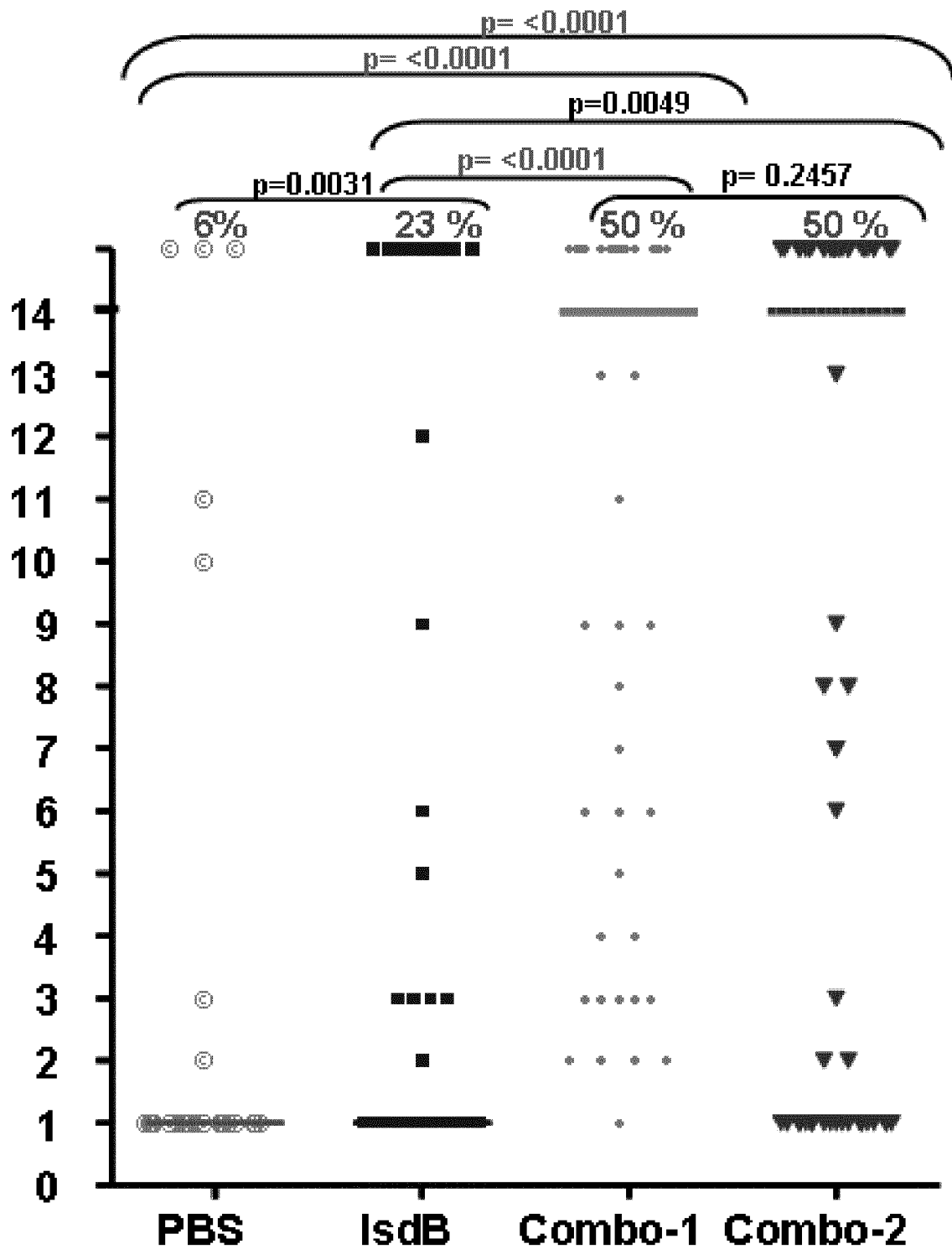


FIGURE 6

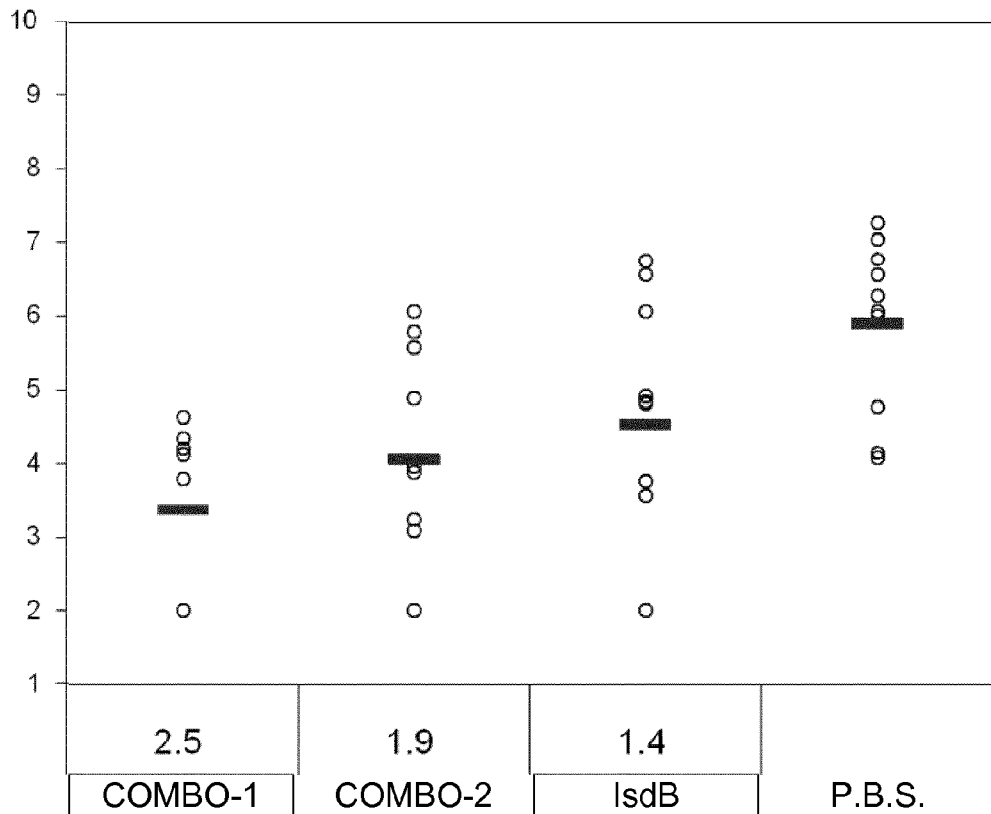


FIGURE 8

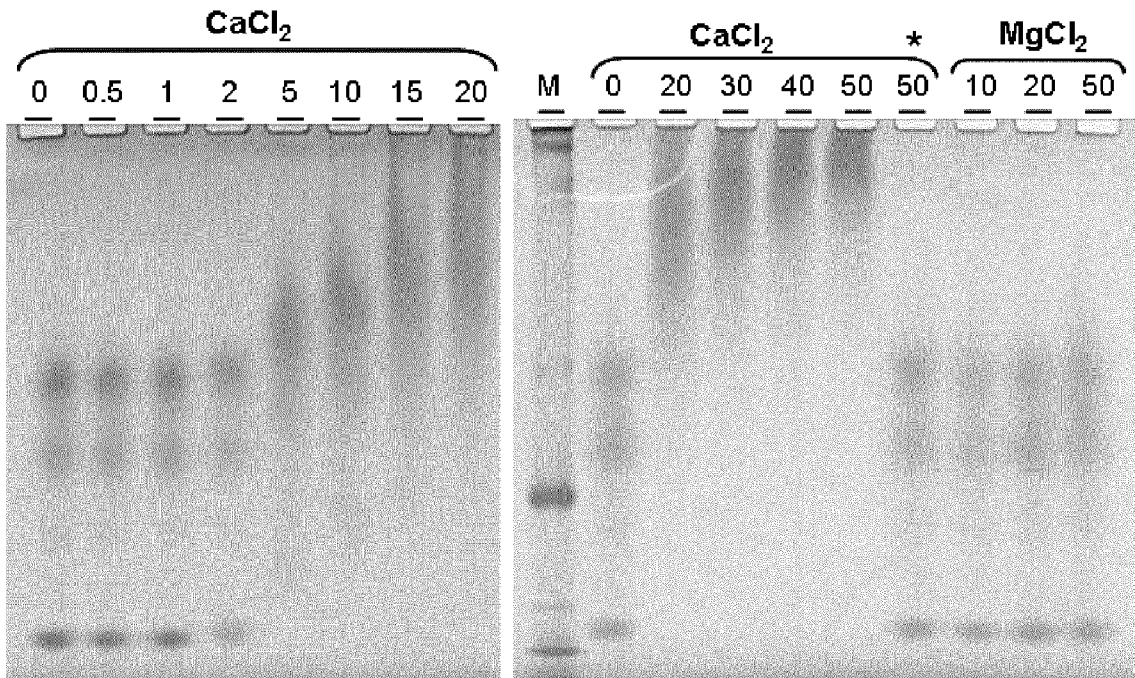


FIGURE 7

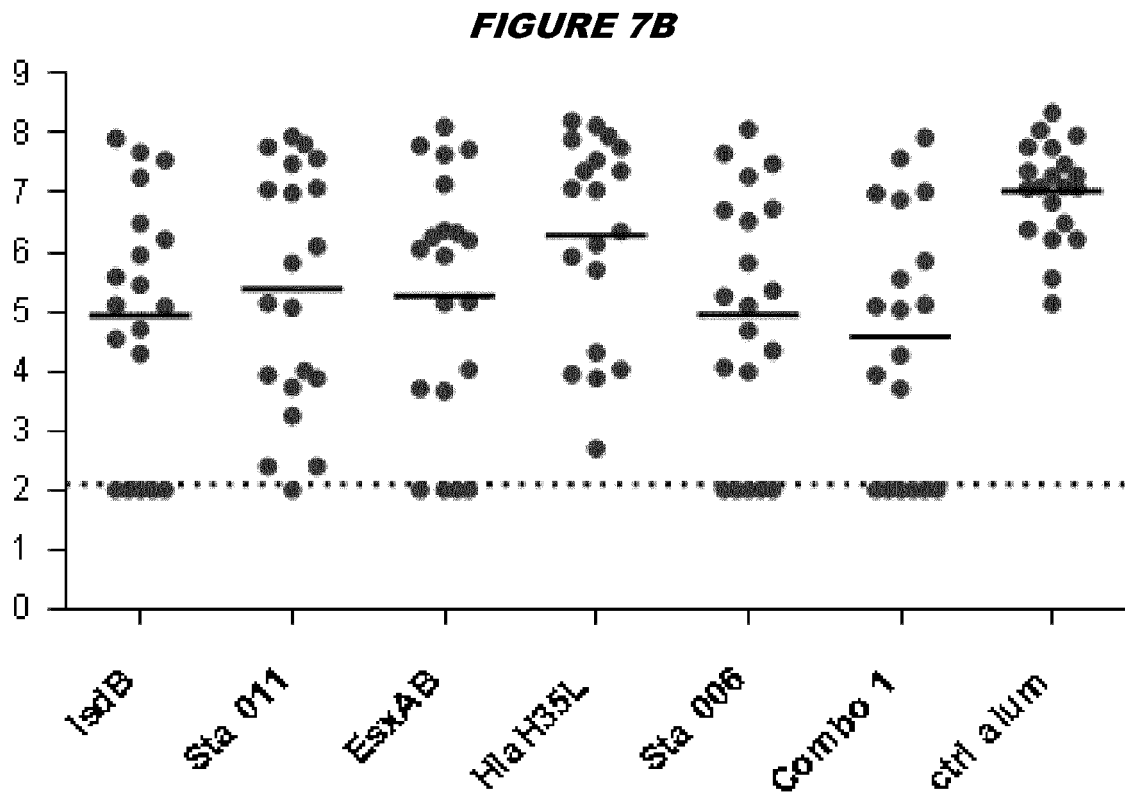
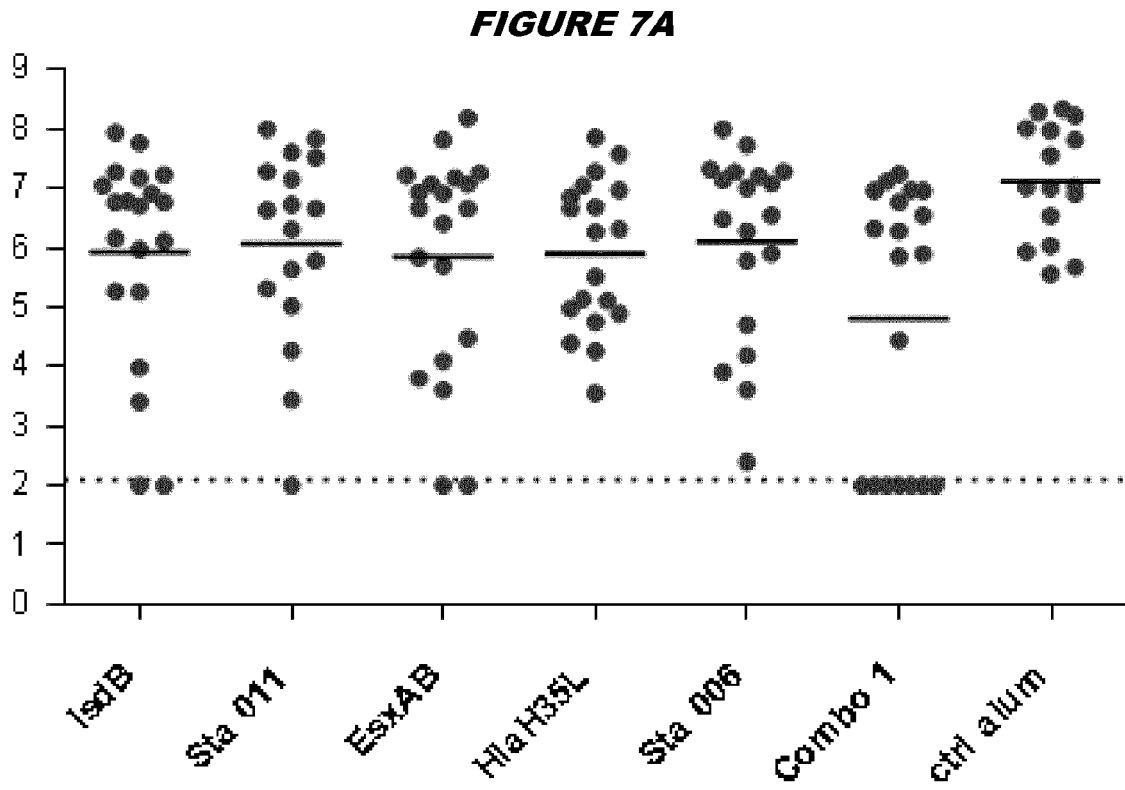


FIGURE 7C

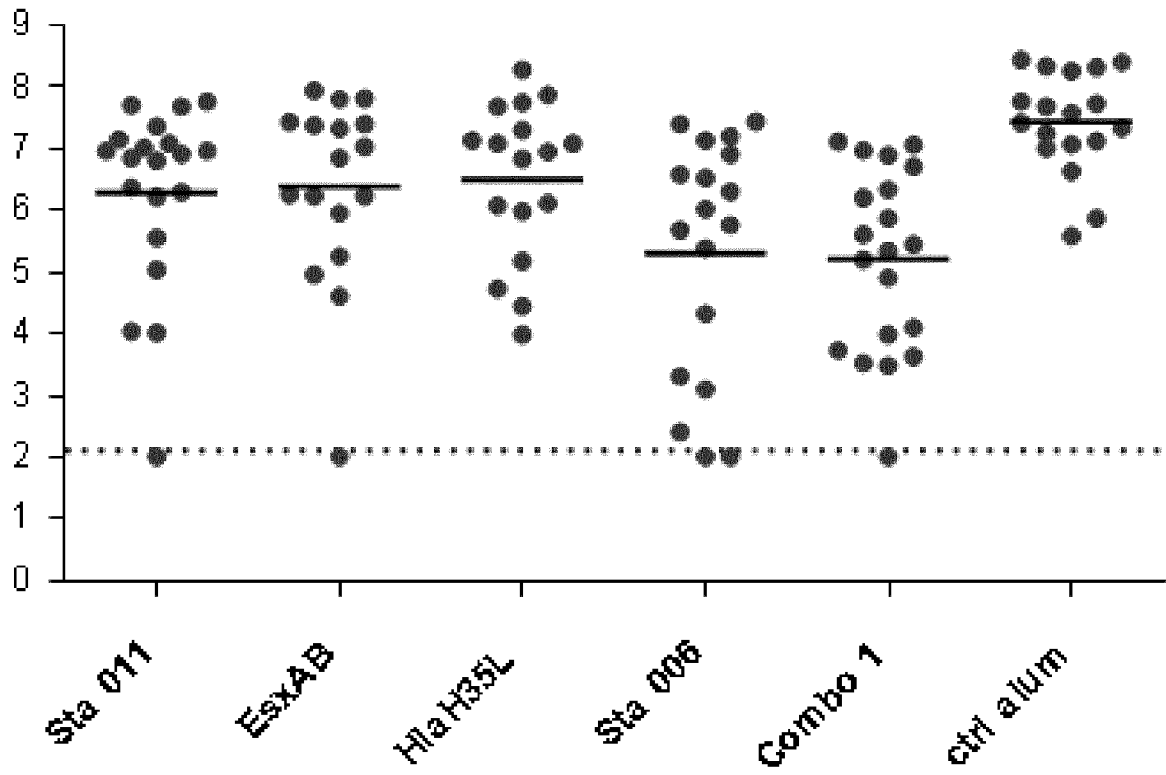


FIGURE 7D

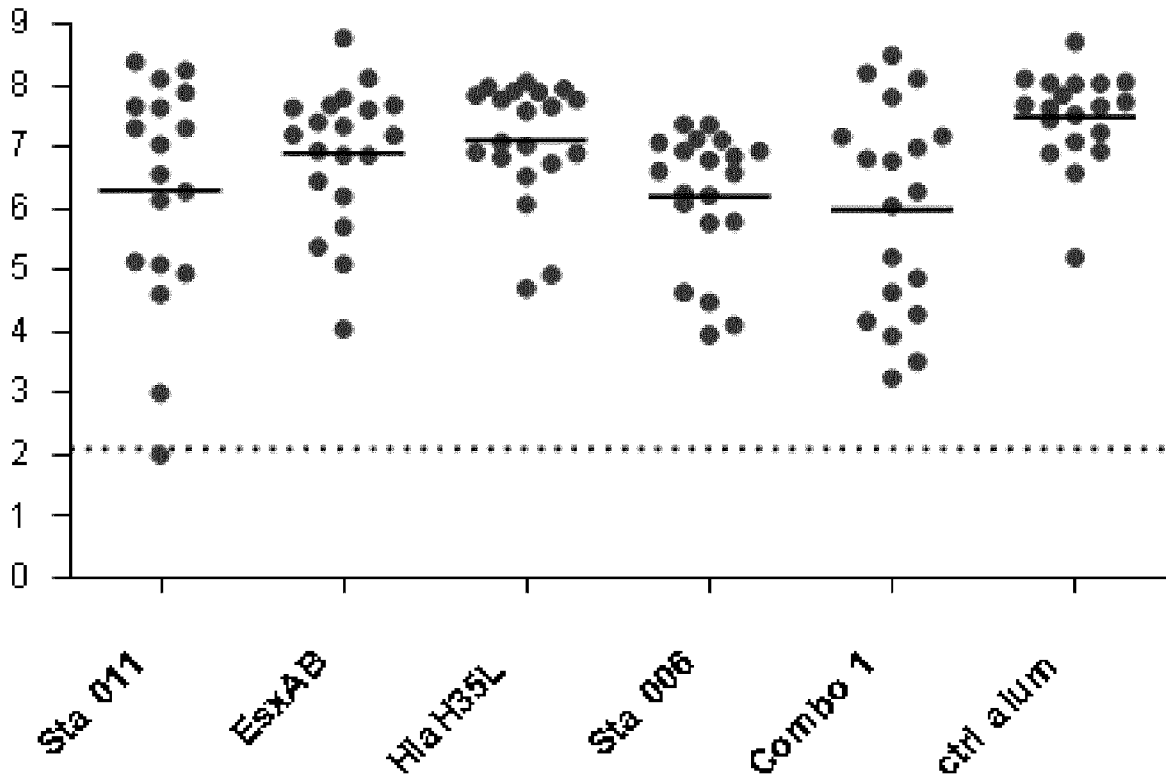


FIGURE 9

FIGURE 9A

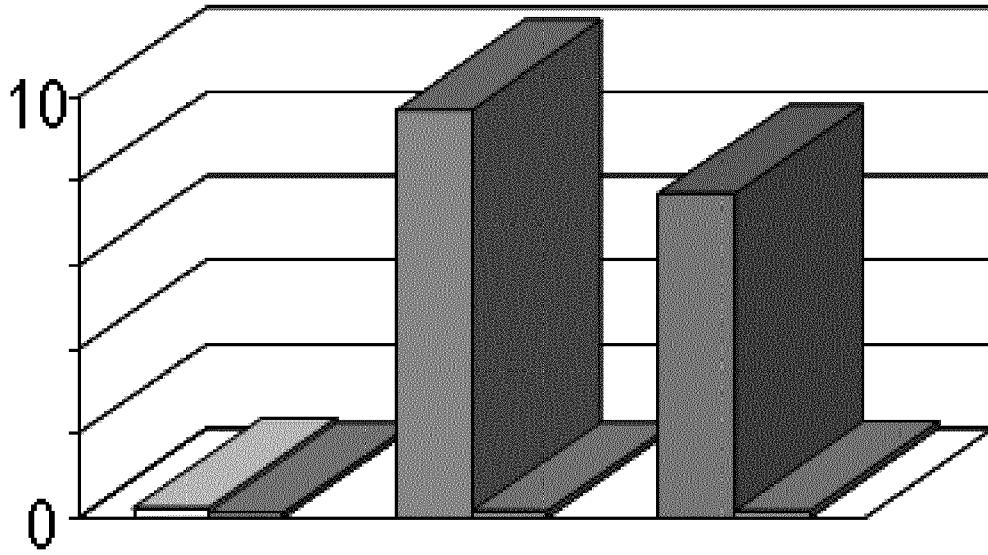


FIGURE 9B

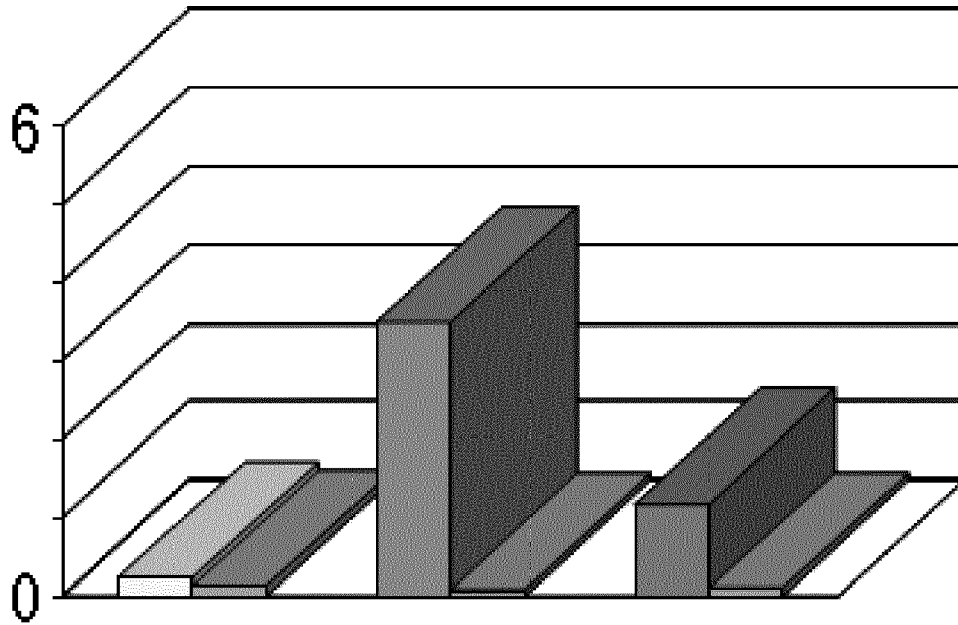


FIGURE 9C

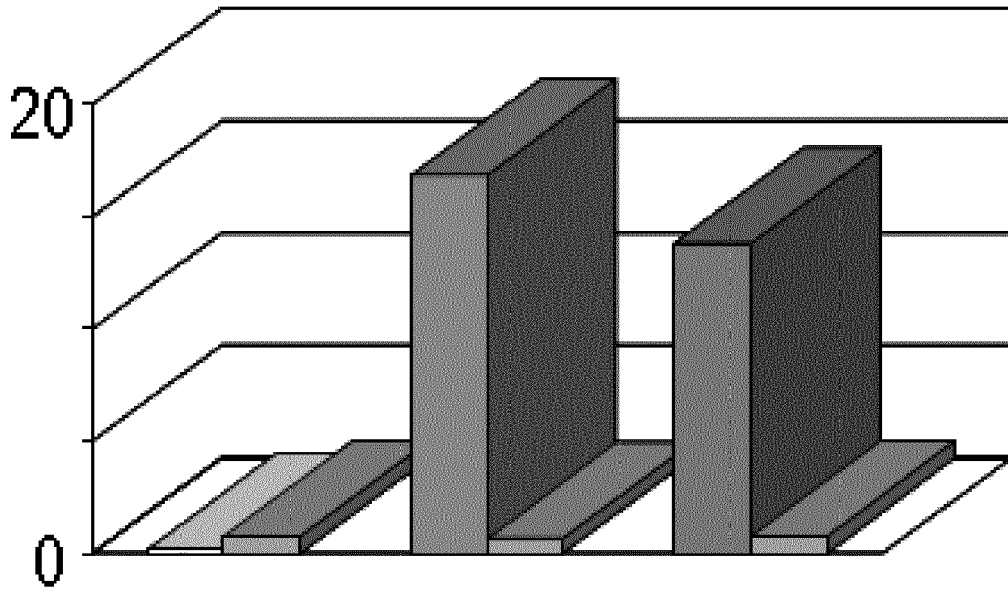


FIGURE 9D

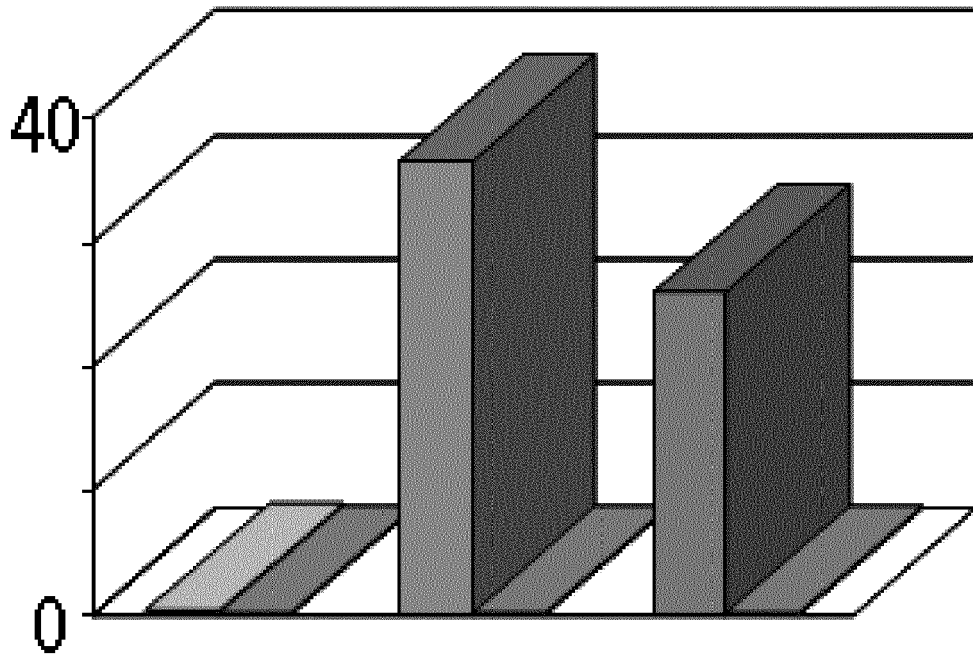


FIGURE 10

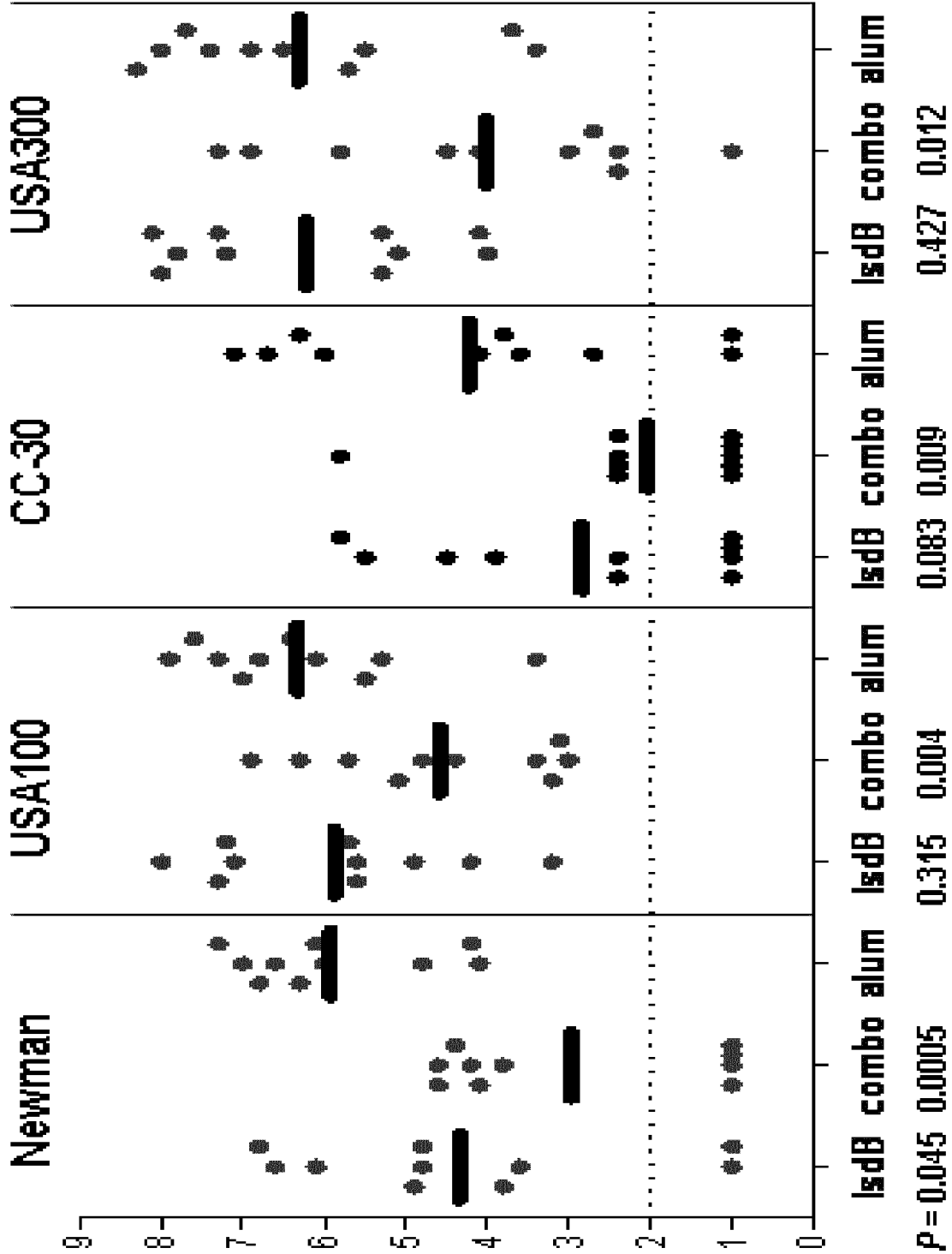


FIGURE 11

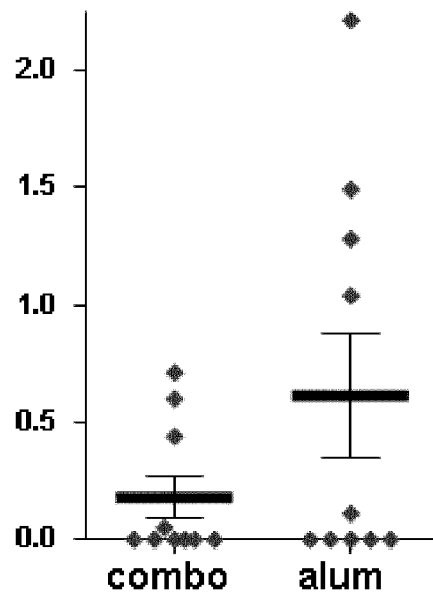


FIGURE 12

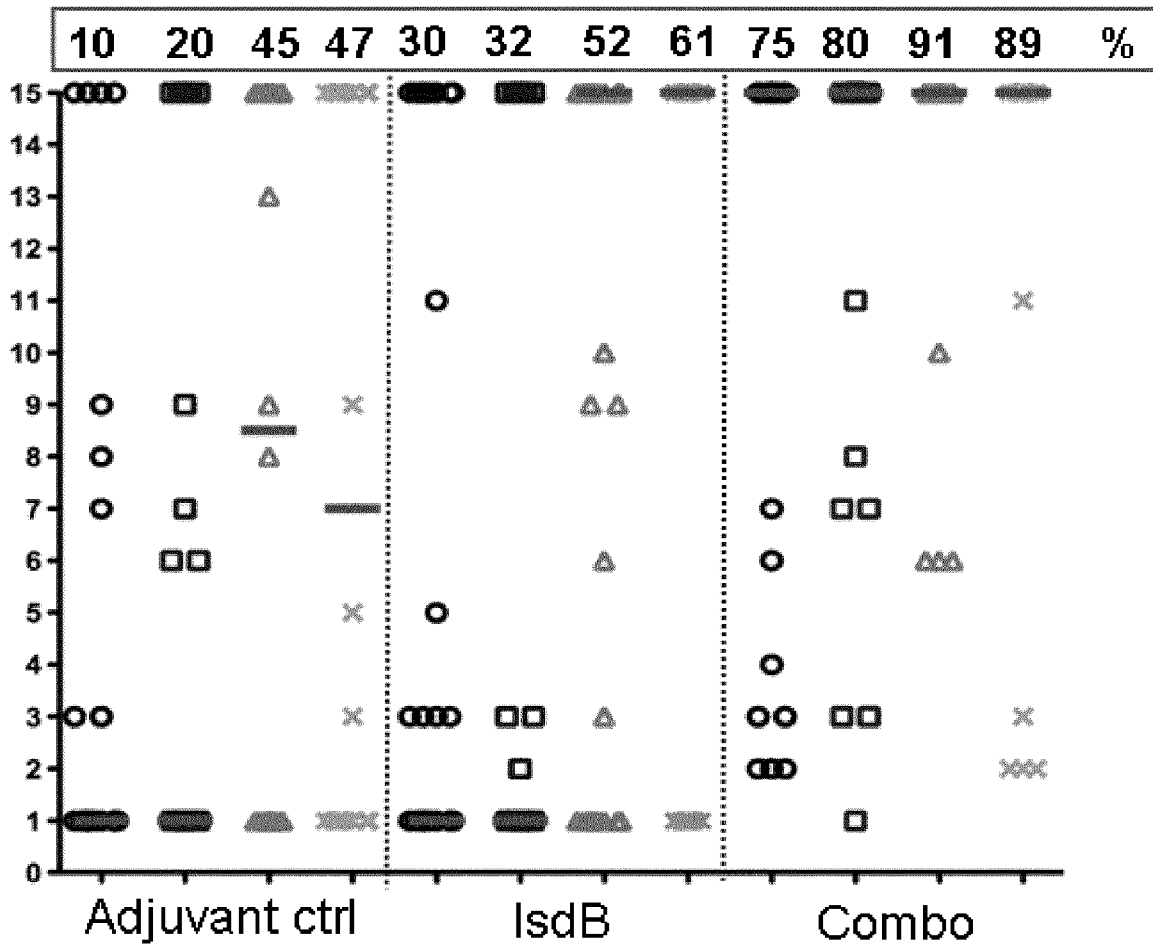
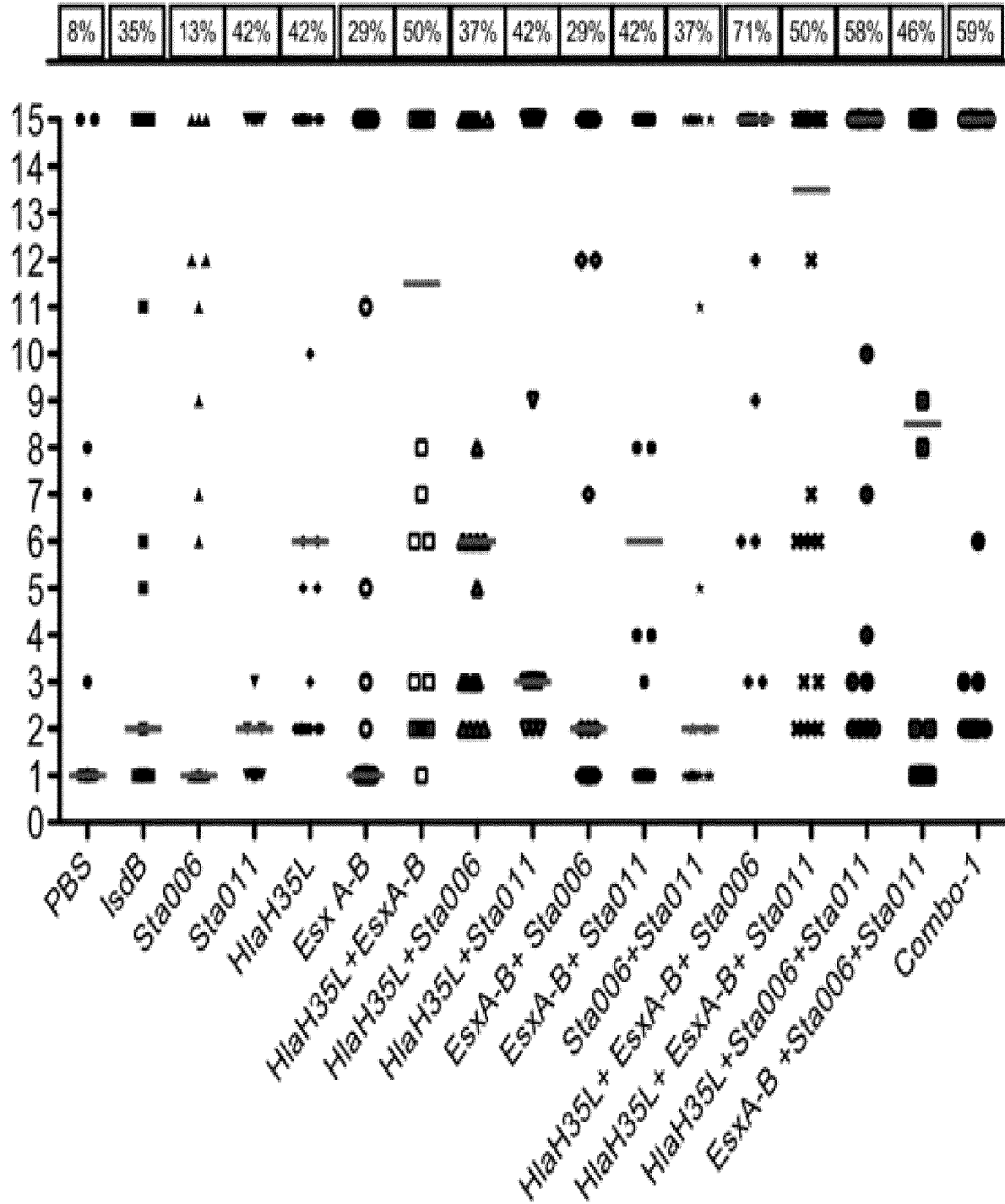


FIGURE 13





EUROPEAN SEARCH REPORT

 Application Number
 EP 12 17 5868

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Category	Citation of document with indication, where appropriate, of relevant passages	Relevant to claim	CLASSIFICATION OF THE APPLICATION (IPC)
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A	SCHAFFER ADAM C ET AL: "Vaccination and passive immunisation against Staphylococcus aureus.", INTERNATIONAL JOURNAL OF ANTIMICROBIAL AGENTS NOV 2008 LNKD- PUBMED:18757184, vol. 32 Suppl 1, November 2008 (2008-11), pages S71-S78, XP002601981, ISSN: 0924-8579 * e.g. section 6.2 on page S75; page S77, right-hand column, paragraph 2; the whole document * -----	1-15	
A	DATABASE Geneseq [Online] 20 November 2003 (2003-11-20), XP002601982, retrieved from EBI accession no. GSP:ABM72354 Database accession no. ABM72354 * the whole document * -----	1-15	TECHNICAL FIELDS SEARCHED (IPC) A61K C07K
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Place of search		Date of completion of the search	Examiner
The Hague		24 August 2012	Gruber, Andreas
CATEGORY OF CITED DOCUMENTS X : particularly relevant if taken alone Y : particularly relevant if combined with another document of the same category A : technological background O : non-written disclosure P : intermediate document T : theory or principle underlying the invention E : earlier patent document, but published on, or after the filing date D : document cited in the application L : document cited for other reasons & : member of the same patent family, corresponding document			

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