



US 20190271019A1

(19) **United States**

(12) **Patent Application Publication**
Silverman et al.

(10) **Pub. No.: US 2019/0271019 A1**

(43) **Pub. Date: Sep. 5, 2019**

(54) **CARBOHYDRATE-ENRICHED
RECOMBINANT MICROORGANISMS**

(71) Applicant: **CALYSTA, INC.**, Menlo Park, CA
(US)

(72) Inventors: **Joshua A. Silverman**, Los Altos Hills,
CA (US); **Lorraine Joan Giver**,
Sunnyvale, CA (US); **Jana Mueller**,
Sunnyvale, CA (US); **Renee M. Saville**,
Mountain View, CA (US); **Drew D.
Regitsky**, San Francisco, CA (US)

(21) Appl. No.: **16/413,422**

(22) Filed: **May 15, 2019**

Related U.S. Application Data

(62) Division of application No. 15/111,733, filed on Jul.
14, 2016, now Pat. No. 10,337,043, filed as applica-
tion No. PCT/US2015/011860 on Jan. 16, 2015.

(60) Provisional application No. 61/928,366, filed on Jan.
16, 2014.

Publication Classification

(51) **Int. Cl.**

<i>C12P 19/18</i>	(2006.01)
<i>C12N 9/00</i>	(2006.01)
<i>A23K 50/00</i>	(2006.01)
<i>A23K 10/12</i>	(2006.01)
<i>A23K 20/163</i>	(2006.01)
<i>C12P 19/04</i>	(2006.01)
<i>C12N 9/10</i>	(2006.01)

(52) **U.S. Cl.**

CPC *C12P 19/18* (2013.01); *C12N 9/00*
(2013.01); *A23K 50/00* (2016.05); *C12Y*
204/01034 (2013.01); *A23K 20/163* (2016.05);
C12P 19/04 (2013.01); *C12N 9/1051*
(2013.01); *A23K 10/12* (2016.05)

(57)

ABSTRACT

The present disclosure relates to recombinant microorgan-
isms engineered for enhanced production of a desired car-
bohydrate, as well as related biomass, and compositions
which are useful, inter alia, as animal feed ingredients. The
present disclosure also provides related methods.

Specification includes a Sequence Listing.

CARBOHYDRATE-ENRICHED RECOMBINANT MICROORGANISMS

STATEMENT REGARDING SEQUENCE LISTING

[0001] The “Sequence Listing” submitted electronically concurrently herewith pursuant 37 C.F.R. § 1.821 in computer readable form (CRF) via EFS-Web as file name 200206_416D1_SEQUENCE LISTING.txt is incorporated herein by reference. The electronic copy of the Sequence Listing was created on May 10, 2019, and the size is 277 KB.

TECHNICAL FIELD

[0002] The present disclosure relates to novel recombinant C_1 metabolizing microorganisms comprising an engineered metabolic pathway for the enhanced production of carbohydrates, and related compositions and methods.

BACKGROUND

[0003] Advances in the efficiency in animal feed utilization have been achieved over the past several decades through the use of feed additives. These added substances augment the nutrient-content, energy-content, and/or disease fighting properties of animal feed compositions. A growing challenge for commercial animal producers is the rising cost of grain. The rising costs are due in part to competing demands for grains for biofuel and human food use. With the rising cost of grain and protein components, coupled with limited land available for feed production, alternative low cost animal feed products with beneficial nutritive and disease fighting properties would be highly desirable.

SUMMARY

[0004] In one embodiment, the present disclosure provides a recombinant C_1 metabolizing microorganism comprising an exogenous nucleic acid selected from the group consisting of an exogenous nucleic acid that encodes a carbohydrate biosynthesis enzyme and an exogenous nucleic acid that encodes an expression control sequence that is operably linked to a nucleic acid encoding a native carbohydrate biosynthesis enzyme, wherein the recombinant C_1 metabolizing microorganism is capable of converting a natural gas-derived carbon feedstock into a desired carbohydrate. Typically, the natural gas-derived carbon feedstock is natural gas or methane.

[0005] In another embodiment, the present disclosure provides a biomass derived from the recombinant C_1 metabolizing microorganism of the present disclosure.

[0006] In a further embodiment, the present disclosure provides a carbohydrate composition comprising carbohydrates extracted from the biomass of the present disclosure, wherein the composition exhibits a $\delta^{13}C$ that is less than -30% .

[0007] In a still further embodiment, the present disclosure provides an animal feed comprising the biomass of the present disclosure.

[0008] In another embodiment, the present disclosure provides a culture or fermentation medium comprising the biomass or composition of the present disclosure.

[0009] The present disclosure additionally provides related methods.

DETAILED DESCRIPTION

[0010] The instant disclosure provides novel recombinant C_1 metabolizing microorganisms that have the ability to utilize relatively low-cost carbon feedstock as an energy source, as well as related biomass, compositions, and methods. The recombinant microorganisms of the present disclosure are engineered for the enhanced production of certain carbohydrates that are commercially desirable. These recombinant microorganisms, as well as the biomass and carbohydrate compositions that are derived from them, are useful as a source of nutrition for animals (such as, for example, livestock, fish, poultry, and the like), as well as cultured or fermented microorganisms.

[0011] In one embodiment, the present disclosure provides a recombinant C_1 metabolizing microorganism, wherein the recombinant C_1 metabolizing microorganism comprises an exogenous nucleic acid selected from the group consisting of an exogenous nucleic acid that encodes a carbohydrate biosynthesis enzyme and an exogenous nucleic acid that encodes an expression control sequence that is operably linked to a nucleic acid encoding a native carbohydrate biosynthesis enzyme, wherein the recombinant C_1 metabolizing microorganism is capable of converting a natural gas carbon feedstock into the carbohydrate. When these recombinant microorganisms are cultured in the presence of a natural gas-derived C_1 substrate, they typically exhibit a $\delta^{13}C$ of less than -30% , and often less than -40% , as described in more detail herein. Typically, the recombinant microorganism is a non-photosynthetic C_1 metabolizing microorganism.

[0012] In these embodiments, the recombinant microorganisms of the present disclosure are engineered to convert a natural gas-derived feedstock, which is a relatively low cost and abundant resource (for example, natural gas, or a C_1 substrate such as methane from natural gas) as compared to more costly carbohydrates, to higher valued carbohydrates. As used herein, the term “natural gas-derived feedstock” refers to natural gas, or any of the components isolated from natural gas (including C_1 substrates) or converted from natural gas (i.e., syngas).

[0013] The term “natural gas” refers herein to naturally occurring gas mixtures that may be obtained by conventional processes (e.g., drilling and water flooding of porous reservoirs) or non-conventional processes (e.g., hydraulic fracturing, horizontal drilling or directional drilling of formations having low gas permeability). The gas mixtures are made up of methane and other compounds, including other C_1 compounds, as well as other light alkane gases (such as, for example, ethane, propane, butane, pentane, and the like), carbon dioxide, nitrogen, hydrogen sulfide, or the like, and combinations thereof. Unconventional natural gas may be obtained from sources such as, for example, tight gas sands formed in sandstone or carbonate, coal bed methane formed in coal deposits and adsorbed in coal particles, shale gas formed in fine-grained shale rock and adsorbed in clay particles or held within small pores or microfractures, methane hydrates that are a crystalline combination of natural gas and water formed at low temperature and high pressure in places such as under oceans and permafrost.

[0014] As used herein, “ C_1 substrate” or “ C_1 compound” refers to any carbon containing molecule or composition that lacks a carbon-carbon bond. Exemplary C_1 substrates include syngas, methane, methanol, formaldehyde, formic acid or a salt thereof, carbon monoxide, carbon dioxide,

methylated amines (e.g., methylamine, dimethylamine, trimethylamine, etc.), methylated thiols, methyl halogens (e.g., bromomethane, chloromethane, iodomethane, dichloromethane, etc.), cyanide, or any combination thereof.

[0015] In certain embodiments of the present disclosure, a natural gas-derived feedstock may be natural gas, a C₁ substrate from natural gas, or syngas. Typically, a C₁ substrate is methane. Exemplary recombinant C₁ metabolizing microorganisms that have utilized a natural gas-derived carbon substrate as a feedstock exhibit a distinctive isotopic carbon signature, which is described in more detail herein. This distinctive isotopic carbon signature is also exhibited by the compositions and products of such recombinant microorganisms (e.g., biomass, carbohydrate compositions, and the like).

[0016] In another embodiment, the present disclosure provides a recombinant C₁ metabolizing microorganism comprising an exogenous nucleic acid encoding a carbohydrate biosynthesis enzyme, wherein the C₁ metabolizing microorganism is capable of converting methane into a carbohydrate. Exemplary carbohydrates are glucans. In some embodiments, a carbohydrate is a β -(1,3)-glucan, and may be branched or unbranched or a mixture thereof. Usually, a C₁ metabolizing microorganism is a non-photosynthetic C₁ metabolizing microorganism.

[0017] As used herein, “C₁ metabolizing microorganism” or “C₁ metabolizing non-photosynthetic microorganism” refers to any microorganism having the ability to use a C₁ substrate as a source of energy or as its primary source of energy and biomass, and may or may not use other carbon substrates (such as sugars and complex carbohydrates) for energy and biomass. For example, a C₁ metabolizing microorganism may oxidize a C₁ substrate, such as methane or methanol. C₁ metabolizing microorganisms include bacteria (such as methanotrophs and methylotrophs) and yeast. In certain embodiments, a C₁ metabolizing microorganism does not include a photosynthetic microorganism, such as algae. In some embodiments, the C₁ metabolizing microorganism will be an “obligate C₁ metabolizing microorganism,” meaning its sole source of energy are C₁ substrates. In further embodiments, a C₁ metabolizing microorganism (e.g., methanotroph) will be cultured in the presence of a C₁ substrate feedstock (i.e., using the C₁ substrate as a source of energy).

[0018] Recombinant C₁ metabolizing microorganisms of the present disclosure are engineered for enhanced production of a desired carbohydrate and in one embodiment, comprise an exogenous nucleic acid encoding a carbohydrate biosynthesis (CB) enzyme. The terms “carbohydrate biosynthesis enzyme” and “CB enzyme” are used interchangeably herein to refer to an enzyme that is involved in the production of a carbohydrate by the recombinant host C₁ metabolizing microorganism.

[0019] Exogenous nucleic acids encoding CB enzymes that are employed in the practice of the present disclosure are typically codon optimized for optimal expression from the recombinant host C₁ metabolizing microorganism and encode an enzyme that is either native to a species heterologous to the host C₁ microorganism or is a mutant (i.e., variant) of an enzyme that exists in nature.

[0020] As used herein, the term “carbohydrate” refers to a monosaccharide, a disaccharide, or a polysaccharide. Suitable exogenous nucleic acids employed in the practice of the present disclosure include those which encode enzymes that

are involved in the production of a monosaccharide such as, for example, glucose, fructose, ribose, glyceraldehyde, galactose and the like; a disaccharide, such as, for example lactose, sucrose, maltose, cellulobiose, and the like, and mixtures thereof or a polysaccharide, including, for example, an unbranched or branched glucan, and the like, and mixtures thereof. Exemplary glucans include α -glucans, such as for example, dextran, glycogen, pullulan, starch, and the like, as well as β -glucans, such as, for example, β -1,4-glucan (i.e., cellulose), β -1,3-glucan, β -(1,3)(1,4)-glucan, β -(1,3)(1,6)-glucan, and the like, and mixtures thereof.

[0021] In a specific embodiment, the CB enzyme is an enzyme involved in the production of an unbranched or a branched glucan, or mixture thereof β -glucans are known to have beneficial therapeutic properties, including as a powerful immune stimulant and a powerful antagonist to both benign and malignant tumors. β -glucans are also known to lower cholesterol and triglyceride levels. See D. Akramiené et al., *Medicina (kaunas)*, 2007; 43(8):597. The β -glucans are a heterogeneous group of glucose polymers made up of β -D-glucopyranosyl units having β -(1,3) and/or β -(1,4), and/or β -(1,6) linkages. They have been isolated from a number of sources, including plants (oat, barley, bran, seaweed, corn, soy, and the like), bacteria (e.g., *Pneumocystis carinii*, *Cryptococcus neoformans*, *Aspergillus fumigatus*, *Histoplasma capsulatum*, *Candida albicans*, and the like), and fungi (i.e., *Saccharomyces cerevisiae* and mushrooms, such as, for example shiitake (*Lentinus edodes*), maitake (*Grifola frondosa*), schizophylan (*Schizophillum commune*), and SSG (*Sclerotinia sclerotiorum*). β -glucan extracts from *Lentinus edodes* and *Schizophillum commune* have been used for the treatment of cancer in Japan since 1980. Id.

[0022] Exogenous nucleic acids that are suitable for use in the practice of the present disclosure include those which encode enzymes involved in gluconeogenesis, glycogenesis, α - or β -glucan biosynthesis, and other metabolic pathways known to produce a carbohydrate.

[0023] Suitable exogenous nucleic acids include those which encode a gluconeogenesis enzyme selected from the group consisting of a pyruvate carboxylase, a phosphoenolpyruvate carboxykinase, an enolase, a phosphoglycerate mutase, a phosphoglycerate kinase, a glyceraldehyde-3-phosphate dehydrogenase, a Type A aldolase, a fructose 1,6-bisphosphatase, a phosphofructokinase, a phosphoglucose isomerase, a hexokinase, a glucose-6-phosphate, and the like.

[0024] Other suitable exogenous nucleic acids include those which encode a glycogenesis enzyme selected from the group consisting of a glucose-1-phosphate adenylyltransferase, a glycogen synthase, and the like.

[0025] The above enzymes can be found in a number of heterologous species, including microorganisms, such as, for example, bacteria and yeast, including, for example, *E. coli*, *C. glutamicum*, *Saccharomyces cerevisiae*, and the like, as well as higher order fungi, such as mushrooms, and the like, as well as algae, and plants.

[0026] Suitable exogenous nucleic acids include those which encode a glucan biosynthesis enzyme, such as, for example, a glucan synthase. An exemplary glucan synthase is β -1,3-glucan synthase. The exogenous nucleic acid may encode a glucan biosynthesis enzyme (e.g., a glucan synthase (such as, for example a β -1,3-glucan synthase)) from a plant (oat, barley, bran, seaweed, corn, soy, and the like),

a bacteria (e.g., *Pneumocystis carinii*, *Cryptococcus neoformans*, *Aspergillus fumigatus*, *Histoplasma capsulatum*, *Candida albicans*, and the like), or a fungi (i.e., *Saccharomyces cerevisiae* and mushrooms, such as, for example shiitake (*Lentinus edodes*), maitake (*Grifola frondosa*), schizophylan (*Schizophyllum commune*), and SSG (*Sclerotinia sclerotiorum*). The amino acid and nucleic acid sequences of a number of β -(1,3)-glucan synthases are known. See, e.g., U.S. Pat. No. 5,194,600, WO99/49047, and EP 0 724 644 B1, all of which are incorporated herein by reference. In certain specific embodiments, the exogenous nucleic acid encodes a carbohydrate biosynthesis enzyme having the amino acid sequence of any of SEQ NOs: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, or 38, shown in Table A, hereinbelow. As described above, the exogenous nucleic acid is typically codon optimized for optimal expression from the recombinant C_1 microorganism. Exemplary nucleic acid sequences encoding these CB enzymes are also provided in Table A. These nucleic acid sequences have been codon optimized for expression in *Methylococcus capsulatus* Bath.

TABLE A

Exemplary Carbohydrate Biosynthesis Enzymes		
Source/Enzyme Name	Amino Acid Sequence (SEQ ID NO.)	Nucleic Acid Sequence (SEQ ID NO.)
<i>Saccharomyces cerevisiae</i> : mature KRE1 protein	2	1
<i>Saccharomyces cerevisiae</i> : mature KRE2 protein	4	3
<i>Saccharomyces cerevisiae</i> s288c: FKS1	6	5
<i>Saccharomyces cerevisiae</i> : FKS2	8	7
<i>Candida albicans</i> : FKS1	10	9
<i>Zea mays</i> (corn): portion of 1,3- β -D-glucan synthase	12	11
<i>Zea mays</i> (corn): portion of 1,3- β -D-glucan synthase	14	13
<i>Oryza sativa</i> (rice): portion of 1,3-beta-D-glucan synthase	16	15
<i>Oryza sativa</i> (rice): portion of 1,3-beta-D-glucan synthase	18	17
<i>Glycine max</i> (soy): portion of 1,3-beta-D-glucan synthase.	20	19
<i>Veronia mespilifolia</i> : 1,3-beta-D-glucan synthase	22	21
<i>Triticum aestivum</i> (wheat): 1,3-beta-D-glucan synthase	24	23
<i>Hordeum vulgare</i> (barley): 1,3-beta-D-glucan synthase	26	25
<i>E. coli</i> : Glucose-1-phosphate adenylyltransferase (Acc. No. YP 49003.1)	28	27
<i>Cornibacterium. Glutamicum</i> (ATCC 13032): Glucose-1-phosphate adenylyltransferase	30	29
<i>Escherichia coli</i> str. K-12 substr. W3110: Glycogen Synthase	32	31
<i>Cornibacterium glutamicum</i> (ATCC 13032): Glycosyltransferase	34	33
<i>E. coli</i> : 1,4-alpha-glucan branching enzyme (Acc. No. YP 492001.1)	36	35
<i>Corynebacterium glutamicum</i> (ATCC 13032): Glycogen branching enzyme	38	37

[0027] Suitable exogenous nucleic acids employed in the practice of the present disclosure include those which encode a variant CB enzyme sequence that is at least 90%, at least 91%, at least 92%, at least 93%, at least 94%, at least 95%, at least 96%, at least 97%, at least 98%, or at least 99% identical to a reference or parental wild-type polypeptide sequence, such as, for example a reference sequence corresponding to any one of SEQ ID NOs: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, or 38, provided that the variant retains the carbohydrate biosynthesis enzyme activity of interest. In certain embodiments, the CB enzyme variant polypeptides will include at least one amino acid substitution (e.g., 1, 2, 3, 5, 6, 7, 8, 9 or 10 or more or up to 20, 25, or 30 substitutions) at a pre-determined position relative to a reference or parental wild-type CB enzyme, provided that a variant retains the CB enzyme activity of interest. The CB enzyme variant polypeptides may further comprise one or more conservative substitutions. A "conservative substitution" is recognized in the art as a substitution of one amino acid for another amino acid that has similar properties. Exemplary conservative substitutions are well known in the art (see, e.g., WO 97/09433, p. 10; Lehninger, Biochemistry, 2nd Edition; Worth Publishers, Inc. NY:NY (1975), pp. 71-77; Lewin, Genes IV, Oxford University Press, NY and Cell Press, Cambridge, Mass. (1990), p. 8, which are incorporated herein by reference). Methods for generating suitable exogenous nucleic acids encoding such variant enzymes are described in more detail herein.

[0028] The "percent identity" between two or more nucleic acid or amino acid sequences is a function of the number of identical positions shared by the sequences (i.e., % identity=number of identical positions/total number of positions \times 100), taking into account the number of gaps, and the length of each gap that needs to be introduced to optimize alignment of two or more sequences. The comparison of sequences and determination of percent identity between two or more sequences can be accomplished using a mathematical algorithm, such as BLAST and Gapped BLAST programs at their default parameters (e.g., Altschul et al., *J. Mol. Biol.* 215:403, 1990; see also BLASTN at the world wide web at ncbi.nlm.nih.gov/BLAST, which are incorporated herein by reference).

[0029] As indicated above, the exogenous nucleic acids encoding CB enzymes employed in the practice of the present disclosure may be codon optimized for expression in the C_1 metabolizing microorganism. Expression of recombinant proteins may be difficult outside their original host. For example, variation in codon usage bias has been observed across different species of bacteria (Sharp et al., *Nucl. Acids. Res.* 33:1141, 2005, which is incorporated herein by reference). Overexpression of recombinant proteins even within their native host may also be difficult. In certain embodiments, the nucleic acid to be introduced into a host as described herein may be subjected to codon optimization prior to introduction into the host to ensure protein expression is effective or enhanced. Codon optimization refers to alteration of codons in genes or coding regions of nucleic acids before transformation to reflect the typical codon usage of the host without altering the polypeptide encoded by the non-natural DNA molecule. Codon optimization methods for optimum gene expression in heterologous hosts have been previously described (see, e.g., Welch et al., *PLoS One* 4:e7002, 2009; Gustafsson et al., *Trends Biotechnol.* 22:346, 2004; Wu et al., *Nucl. Acids Res.*

35:D76, 2007; Villalobos et al., *BMC Bioinformatics* 7:285, 2006; U.S. Patent Publication Nos. 2011/0111413 and 2008/0292918; disclosure of which methods are incorporated herein by reference, in their entirety). Exogenous nucleic acids encoding CB enzymes that are suitable for use in the practice of the present disclosure include those having a nucleic acid sequence that is at least about 85% identical to a nucleic acid reference sequence selected from the group consisting of SEQ ID NO.:1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, and 37. In some embodiments, the exogenous nucleic acid encoding the CB enzyme has a nucleic acid sequence that is at least about 85%, at least about 86%, at least about 87%, at least about 88%, at least about 89%, at least about 90%, at least about 91%, at least about 92%, at least about 93%, at least about 94%, at least about 95%, at least about 96%, at least about 97%, at least about 98% and at least about 99% sequence identity to a nucleic acid reference sequence selected from the group consisting of SEQ ID NO.:1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, and 37. Illustrative exogenous nucleic acids that encode a CB enzyme which are suitable for use in the practice of the invention include sequences which have been codon optimized for optimal expression in *Methylococcus capsulatus* Bath, such as, for example, any one of SEQ ID NO.:1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, and 37.

[0030] Similarly, exogenous nucleic acid molecules of this disclosure encoding polypeptide variants may be designed using the phylogenetic-based methods described in the references noted above (U.S. Pat. No. 8,005,620; Gustafsson et al.; Welch et al.; Villalobos et al.; Minshull et al., all of which are incorporated herein by reference.).

[0031] An exogenous nucleic acid encoding a carbohydrate biosynthesis enzyme includes polynucleotides that encode a polypeptide, a polypeptide fragment, a peptide, or a fusion polypeptide that has or retains the corresponding carbohydrate biosynthesis enzyme activity. Methods to determine whether a polypeptide has a particular activity by measuring the ability of the polypeptide to convert a substrate into a product are known in the art.

[0032] In some embodiments, the exogenous nucleic acid encodes an expression control sequence that is operably linked to a nucleic acid encoding a native carbohydrate biosynthesis enzyme. Typically, the expression control sequence is one that results in the overexpression of a native carbohydrate biosynthesis enzyme. As used herein, “over-expressed” and “overexpression” when referring to a gene or a protein means an increase in expression or activity of the gene or protein. Increased expression or activity includes expression or activity of a gene or protein being increased above the level of a wildtype (native or non-genetically engineered) control or reference microorganism. A gene or protein is overexpressed if the expression or activity is in a microorganism where it is not normally expressed or active. A gene or protein is overexpressed if the expression or activity is extended or present longer in the recombinant microorganism than in a wild-type control or reference microorganism.

[0033] In addition to the exogenous nucleic acids described hereinabove, recombinant C_1 metabolizing microorganisms of the present disclosure may comprise further genetic modifications which enhance the production of the desired carbohydrate. For example, when the exogenous nucleic acid encodes a carbohydrate biosynthesis enzyme,

the recombinant C_1 metabolizing microorganism may further comprise an exogenous expression control sequence that is operatively linked to the exogenous nucleic acid encoding the carbohydrate biosynthesis enzyme to enhance production of the desired carbohydrate. Expression control sequences suitable for use in the practice of the present disclosure are described in more detail herein.

[0034] Alternatively, or in addition, the recombinant C_1 metabolizing microorganism of the present disclosure may further comprise an exogenous expression control sequence operatively linked to an endogenous nucleic acid encoding an endogenous enzyme that utilizes one or more of the same substrates utilized by carbohydrate biosynthesis enzymes, or utilizes the desired carbohydrate as a substrate (i.e., a “competing” endogenous enzyme). This may be done to downregulate the competing endogenous enzyme.

[0035] In some embodiments, it may be desirable to reduce or inhibit a competing endogenous enzyme activity by mutating the competing endogenous enzyme to delete or attenuate its activity. “Inhibit” or “inhibited,” as used herein, refers to an alteration, reduction, down regulation, abrogation or deletion, directly or indirectly, in the expression of a target gene or in the activity of a target molecule relative to a control, endogenous or reference molecule, wherein the alteration, reduction, down regulation or abrogation is statistically, biologically, industrially, or clinically significant.

[0036] Various methods for downregulating, inactivating, knocking-out, or deleting endogenous gene function in C_1 metabolizing microorganisms are known in the art. For example, targeted gene disruption is an effective method for gene down-regulation where an exogenous polynucleotide is inserted into a structural gene to disrupt transcription. Genetic cassettes comprising the exogenous insertion DNA (e.g., a genetic marker) flanked by sequence having a high degree of homology to a portion of the target host gene to be disrupted are introduced into the host C_1 metabolizing microorganism. Exogenous DNA disrupts the target host gene via native DNA replication mechanisms. Allelic exchange to construct deletion/insertional mutants in C_1 metabolizing microorganisms, including methanotrophic bacteria, have been described in, for example, Toyama and Lidstrom, *Microbiol.* 144:183, 1998; Stoylar et al., *Microbiol.* 145:1235, 1999; Ali et al., *Microbiol.* 152:2931, 2006; Van Dien et al., *Microbiol.* 149:601, 2003; Martin and Murrell, *FEMS Microbiol. Lett.* 127:243, 2006, all of which are incorporated herein by reference.

[0037] For example, in some embodiments of the present disclosure, a recombinant C_1 metabolizing microorganisms may further comprise a deletion of endogenous glycogen synthase activity and/or endogenous phosphoglucomutase activity. Enzymes involved in other pathways, such as an amino acid synthesis pathway, may also be targeted for down regulation to focus metabolic activities of the host microorganism on carbohydrate biosynthesis.

[0038] The recombinant C_1 metabolizing microorganism may thus be engineered to have the ability to produce the desired carbohydrate at enhanced levels. In some of these embodiments, a recombinant C_1 metabolizing microorganism produces the desired carbohydrate at a level that is at least about 10% greater than that produced by the native C_1 metabolizing microorganism and up to about 2-fold, to about 3-, 4-, 5-, 10-, 20-, 30-, 40-, 50-, 60-, 70-, 80-, 90-, 100-, and up to about 500- or about 1000-fold the level produced by a native C_1 metabolizing microorganism, when cultured in

the presence of a natural gas-derived feedstock (e.g., natural gas, methane, and the like) under at least one set of culture conditions. In other embodiments, a recombinant C_1 metabolizing microorganism produces the desired carbohydrate at a level that is from at least about 15%, at least about 20%, at least about 25%, at least about 30%, at least about 35%, at least about 40%, at least about 45%, at least about 50%, at least 55%, at least about 60%, at least about 65%, at least about 70%, at least about 75%, at least about 80%, at least about 85%, at least about 90%, or is at least about 95% greater than that produced by a native C_1 metabolizing microorganism, and up to about 2-fold, to about 3-, 4-, 5-, 10-, 20-, 30-, 40-, 50-, 60-, 70-, 80-, 90-, 100-, to about 500- or about 1000-fold the level produced by the native C_1 metabolizing microorganism, when cultured in the presence of a natural gas-derived feedstock under at least one set of culture conditions. Typically, the enhanced level of production of a desired carbohydrate by a recombinant C_1 metabolizing microorganism of the present invention is at least about 2-fold, 3-, 4-, 5-, 10-, 20-, 30-, 40-, 50-, 60-, 70-, 80-, 90-, or 100-fold that of the native C_1 metabolizing microorganism, when cultured in the presence of a natural gas-derived feedstock under at least one set of culture conditions.

[0039] Recombinant methods for expression of exogenous nucleic acids in microbial organisms are well known in the art. Such methods can be found described in, for example, Sambrook et al., *Molecular Cloning: A Laboratory Manual*, Third Ed., Cold Spring Harbor Laboratory, New York (2001); and Ausubel et al., *Current Protocols in Molecular Biology*, John Wiley and Sons, Baltimore, Md. (1999), all of which are incorporated herein by reference. Genetic modifications to nucleic acid molecules encoding enzymes, or functional fragments thereof, can confer a biochemical or metabolic capability to a recombinant cell that is altered from its naturally occurring state.

[0040] As used herein, the terms “endogenous” and “native” when referring to a nucleic acid, polypeptide, such as an enzyme, compound or activity refers to a nucleic acid, polypeptide, compound or activity that is normally present in a host cell. The term “homologous” or “homolog” refers to a molecule or activity from an exogenous (non-native) source that is the same or similar molecule or activity as that found in or derived from a host cell, species or strain.

[0041] As used herein, the term “exogenous” when referring to a nucleic acid molecule, construct or sequence refers to a nucleic acid molecule or portion of a nucleic acid molecule sequence that is not native to a cell in which it is expressed, a nucleic acid molecule or portion of a nucleic acid molecule native to a host cell that has been altered or mutated, or a nucleic acid molecule with an altered expression as compared to the native expression levels under similar conditions. For example, an exogenous control sequence (e.g., promoter, enhancer) may be used to regulate expression of a gene or a nucleic acid molecule in a way that is different than the gene or a nucleic acid molecule that is normally expressed in nature or culture. In certain embodiments, an exogenous nucleic acid molecule may be homologous to a native host cell gene, but may have an altered expression level or have a different sequence or both. In other embodiments, exogenous nucleic acid molecules may not be endogenous to a host cell or host genome, but instead may have been added to a host cell by conjugation, transformation, transfection, electroporation, or the like, wherein

the added molecule may integrate into the host genome or can exist as extra-chromosomal genetic material (e.g., plasmid or other self-replicating vector).

[0042] In certain embodiments, more than one exogenous nucleic acid molecule can be introduced into a host cell as separate nucleic acid molecules, as a polycistronic nucleic acid molecule, as a single nucleic acid molecule encoding a fusion protein, or any combination thereof, and still be considered as more than one exogenous nucleic acid. For example, a C_1 metabolizing microorganism can be modified to express two or more exogenous nucleic acid molecules, which may be the same or different, that encode one or more carbohydrate biosynthesis enzyme as disclosed herein. In certain embodiments, multiple copies of a carbohydrate biosynthesis enzyme-encoding polynucleotide molecule are introduced into a host cell, which may be two, three, four, five, six, seven, eight, nine, ten or more copies of the same carbohydrate biosynthesis enzyme or different carbohydrate biosynthesis enzyme encoding polynucleotides.

Host Cells and Transformation Methods

[0043] In carrying out the practice of the present invention, the exogenous nucleic acids described hereinabove are transformed into a host cell that is a C_1 metabolizing microorganism. The C_1 metabolizing microorganism employed may be natural, strain adapted (e.g., performing fermentation to select for strains with improved growth rates and increased total biomass yield compared to the parent strain), or recombinantly modified to produce or overexpress the carbohydrate biosynthesis enzyme of interest and/or to have increased growth rates. Typically, the C_1 metabolizing microorganism is a non-photosynthetic C_1 microorganism (e.g., is not an algae or a plant).

[0044] In certain embodiments, the present disclosure employs C_1 metabolizing microorganisms that are prokaryotes or bacteria, such as *Methylomonas*, *Methylobacter*, *Methylococcus*, *Methylosinus*, *Methylocystis*, *Methylomicrobium*, *Methanomonas*, *Methylophilus*, *Methylobacillus*, *Methylobacterium*, *Hyphomicrobium*, *Xanthobacter*, *Bacillus*, *Paracoccus*, *Nocardia*, *Arthrobacter*, *Rhodopseudomonas*, or *Pseudomonas*.

[0045] In further embodiments, the C_1 metabolizing bacteria employed is a methanotroph or a methylotroph. Exemplary methanotrophs include *Methylomonas*, *Methylobacter*, *Methylococcus*, *Methylosinus*, *Methylocystis*, *Methylomicrobium*, *Methanomonas*, *Methylocella*, or a combination thereof. Exemplary methylotrophs include *Methylobacterium extorquens*, *Methylobacterium radiotolerans*, *Methylobacterium populi*, *Methylobacterium chloromethanicum*, *Methylobacterium nodulans*, or a combination thereof. As used herein, the term “methylotrophic bacteria” refers to any bacteria capable of oxidizing any compound in any form (e.g., solid, liquid, gas) that contains at least one carbon and that do not contain carbon-carbon bonds. In certain embodiments, a methylotrophic bacterium may be a methanotroph. For example, “methanotrophic bacteria” refers to any methylotrophic bacteria that have the ability to oxidize methane as a source of carbon and energy, which may be the primary source of carbon and energy. Exemplary methanotrophic bacteria include *Methylomonas*, *Methylobacter*, *Methylococcus*, *Methylosinus*, *Methylocystis*, *Methylomicrobium*, or *Methanomonas*.

[0046] Methanotrophic bacteria are classified into three groups based on their carbon assimilation pathways and

internal membrane structure: type I (gamma proteobacteria), type II (alpha proteobacteria), and type X (gamma proteobacteria). Type I methanotrophs use the ribulose monophosphate (RuMP) pathway for carbon assimilation whereas type II methanotrophs use the serine pathway. Type X methanotrophs use the RuMP pathway but also express low levels of enzymes of the serine pathway. Methanotrophic bacteria employed in the practice of the present invention include obligate methanotrophs, which can only utilize C_1 substrates for carbon and energy sources, and facultative methanotrophs, which naturally have the ability to utilize some multi-carbon substrates as a carbon and energy source.

[0047] Exemplary facultative methanotrophs employed in the practice of the present invention include some species of *Methylocella*, *Methylocystis*, and *Methylocapsa* (e.g., *Methylocella silvestris*, *Methylocella palustris*, *Methylocella tun-drae*, *Methylocystis daltona* strain SB2, *Methylocystis bryophila*, and *Methylocapsa aurea* KYG), *Methylobacterium organophilum* (ATCC 27,886), *Methylidium petroleiphilum*, or high growth variants thereof. Exemplary obligate methanotrophic bacteria useful in the practice of the present invention include *Methylococcus capsulatus* Bath (NCIMB 11132), *Methylomonas* sp. 16a (ATCC PTA 2402), *Methylosinus trichosporium* OB3b (NRRL B-11,196), *Methylosinus sporium* (NRRL B-11,197), *Methylocystis parvus* (NRRL B-11,198), *Methylomonas methanica* (NRRL B-11,199), *Methylomonas albus* (NRRL B-11,200), *Methylobacter capsulatus* Y (NRRL B-11,201), *Methylomonas flagellata* sp. AJ-3670 (FERM P-2400), *Methylacidiphilum infernorum*, *Methylacidiphilum fumariolicum*, *Methylomicrobium alcaliphilum*, *Methyloacidia kamchatkensis*, or high growth variants thereof.

[0048] Suitable C_1 metabolizing microorganisms useful in the practice of the present invention include syngas metabolizing bacteria such as, for example, *Clostridium*, *Moorella*, *Pyrococcus*, *Eubacterium*, *Desulfobacterium*, *Carboxydothermus*, *Acetogenium*, *Acetobacterium*, *Acetoanaerobium*, *Butyribacterium*, *Peptostreptococcus*, and the like. Exemplary syngas metabolizing bacteria include *Clostridium autoethanogenum*, *Clostridium ljungdahli*, *Clostridium ragsdalei*, *Clostridium carboxydivorans*, *Butyribacterium methylotrophicum*, *Clostridium woodii*, *Clostridium neopropanoligen*, and the like.

[0049] Other suitable C_1 metabolizing microorganisms useful in the practice of the present invention include eukaryotes such as, for example, yeast, including *Candida*, *Yarrowia*, *Hansenula*, *Pichia*, *Torulopsis*, *Rhodotorula*, and the like.

[0050] Each of the microorganisms of this disclosure may be grown as an isolated culture, with a heterologous organism that may aid with growth, or one or more of these bacteria may be combined to generate a mixed culture. The term "heterologous" when referring to an organism refers to a species that is different from the host cell. In still further embodiments, C_1 metabolizing non-photosynthetic microorganisms of this disclosure are obligate C_1 metabolizing non-photosynthetic microorganisms, such as an obligate methanotroph or methylotroph.

[0051] Any one of the aforementioned C_1 metabolizing microorganisms can be used as a parent or reference host cell to make a recombinant C_1 metabolizing microorganisms of this disclosure. As used herein, "recombinant" refers to a non-naturally-occurring organism, microorganism, cell, nucleic acid molecule, or vector that has at least one genetic

alteration or has been modified by the introduction of an exogenous nucleic acid molecule, or refers to a cell that has been altered such that the expression of an endogenous nucleic acid molecule or gene can be controlled. Recombinant also refers to a cell that is derived from a cell or is progeny of a cell having one or more such modifications. Genetic alterations include, for example, modifications introducing expressible nucleic acid molecules encoding proteins or enzymes, or other nucleic acid molecule additions, deletions, substitutions or other functional alteration of a cell's genetic material. For example, recombinant cells may express genes or other nucleic acid molecules that are not found in identical form within the native cell (i.e., unmodified or wild type cell), or may provide an altered expression pattern of endogenous genes, such genes that may otherwise be over-expressed, under-expressed, minimally expressed, or not expressed at all.

[0052] Any of the recombinant C_1 metabolizing microorganisms or methanotrophic bacteria described herein may be transformed to comprise at least one exogenous nucleic acid to provide the host with a new or enhanced activity (e.g., enzymatic activity) or may be genetically modified to remove or substantially reduce an endogenous gene function using any of a variety of methods known in the art.

[0053] Transformation refers to the introduction of a nucleic acid molecule (e.g., exogenous nucleic acid molecule) into a host cell. The transformed host cell may carry the exogenous nucleic acid molecule extra-chromosomally or integrated in the chromosome. Integration into a host cell genome and self-replicating vectors generally result in genetically stable inheritance of the transformed nucleic acid molecule. Host cells containing the transformed nucleic acid molecules are referred to as "non-naturally occurring" or "genetically engineered" or "recombinant" or "transformed" or "transgenic" cells (e.g., bacteria).

[0054] Expression systems and expression vectors useful for the expression of exogenous nucleic acids in C_1 metabolizing microorganisms (e.g., methanotrophic bacteria) are known.

[0055] Electroporation of C_1 metabolizing bacteria is described herein and has been previously described in, for example, Toyama et al., *FEMS Microbiol. Lett.* 166:1, 1998; Kim and Wood, *Appl. Microbiol. Biotechnol.* 48:105, 1997; Yoshida et al., *Biotechnol. Lett.* 23:787, 2001, and U.S. Patent Appl. Pub. No. 2008/0026005.

[0056] Bacterial conjugation, which refers to a particular type of transformation involving direct contact of donor and recipient cells, is more frequently used for the transfer of nucleic acid molecules into C_1 metabolizing bacteria. Bacterial conjugation involves mixing "donor" and "recipient" cells together in close contact with each other. Conjugation occurs by formation of cytoplasmic connections between donor and recipient bacteria, with unidirectional transfer of newly synthesized donor nucleic acid molecules into the recipient cells. A recipient in a conjugation reaction is any cell that can accept nucleic acids through horizontal transfer from a donor bacterium. A donor in a conjugation reaction is a bacterium that contains a conjugative plasmid, conjugative transposon, or mobilized plasmid. The physical transfer of the donor plasmid can occur through a self-transmissible plasmid or with the assistance of a "helper" plasmid. Conjugations involving C_1 metabolizing bacteria is described herein and have been previously described in Stolyar et al., *Mikrobiologiya* 64:686, 1995; Motoyama et

al., *Appl. Micro. Biotech.* 42:67, 1994; Lloyd et al., *Arch. Microbiol.* 171:364, 1999; PCT Publication No. WO 02/18617; and Ali et al., *Microbiol.* 152:2931, 2006.

[0057] Expression control sequences suitable for use in the practice of the present invention include, for example, promoters, terminators, enhancers, repressors, inducers, and the like. Promoters suitable for use in the practice of the present invention may be constitutive, leaky, or inducible, and native or non-native to the host cell employed. Exemplary promoters include a pyruvate decarboxylase (PDC) a promoter, a deoxy-xylulose phosphate synthase promoter, a methanol dehydrogenase promoter (MDH) (such as, for example, the promoter in the upstream intergenic region of the *mxoF* gene from *Methylococcus capsulatus* Bath (Acc. No. MCA0779) or the MDH promoter from *M. extorquens* (See Springer et al., *FEMS Microbiol. Lett.* 160:119 (1998)), a hexulose 6-phosphate synthase promoter, a ribosomal protein S16 promoter, a serine hydroxymethyl transferase promoter, a serine-glyoxylate aminotransferase promoter, a phosphoenolpyruvate carboxylase promoter, a T5 promoter, Trc promoter, a promoter for PHA synthesis (Foellner et al., *Appl. Microbiol. Biotechnol.* 40:284, 1993), a pyruvate decarboxylase promoter (Tokuhira et al., *Appl. Biochem. Biotechnol.* 131:795, 2006), the lac operon Plac promoter (Toyama et al., *Microbiol.* 143:595, 1997), a hybrid promoter such as P_{trc} (Brosius et al., *Gene* 27:161, 1984), promoters identified from native plasmid in methylotrophs (EP 296484), methanotrophs, and the like.

[0058] Additionally, suitable homologous or heterologous promoters for high expression of exogenous nucleic acid molecules may be utilized. For example, U.S. Pat. No. 7,098,005 describes the use of promoters for high expression in the presence of methane or methanol of a heterologous coding nucleic acid in C₁ metabolizing bacteria.

[0059] In certain embodiments, regulated expression of exogenous nucleic acids encoding a carbohydrate biosynthesis enzyme may be desirable to optimize growth rate of the non-naturally occurring C₁ metabolizing microorganism and may improve bacterial growth in a variety of carbon source conditions. This may be achieved through the use of an inducible promoter system.

[0060] In certain embodiments, a nucleic acid encoding CB enzyme is operatively linked to an inducible promoter. Inducible promoter systems employed in the practice of the present invention include those known in the art and include tetracycline inducible promoter system; IPTG/lac operon inducible promoter system, heat shock inducible promoter system; metal-responsive promoter systems; nitrate inducible promoter system; light inducible promoter system; ecdysone inducible promoter system, the inducible/regulatable system described for use in methylotrophic and methanotrophic bacteria (see, e.g., U.S. Patent Appl. No. US 2010/0221813, which is incorporated herein by reference), and the like. For example, in one embodiment, the non-naturally occurring C₁ metabolizing microorganism (e.g., methanotroph, methylotroph) comprises: (1) an exogenous nucleic acid encoding CB enzyme, operatively linked to a promoter flanked by lacO operator sequences, and (2) an exogenous nucleic acid encoding a lac repressor protein operatively linked to a constitutive promoter (e.g., hexulose-6-phosphate synthase promoter). Induction is initiated when Lac repressor protein binds to lacO operator sequences flanking the LDH or other promoter, preventing transcription. IPTG binds lac repressor and releases it from lacO

sequences, allowing transcription. By using an inducible promoter system, lactate synthesis may be controlled by the addition of an inducer.

[0061] The expression systems and expression vectors employed in the practice of the present invention optionally contain genetic elements, such as, for example, one or more ribosome binding sites for translation initiation and a transcription termination site, polyadenylation signals, restriction enzyme sites, multiple cloning sites, other coding segments, and the like. In certain embodiments, promoters and/or codon optimization (described in more detail hereinabove) are used for high constitutive expression of exogenous polynucleotides encoding one or more carbohydrate biosynthesis enzymes in host methanotrophic bacteria. Regulated expression of an exogenous nucleic acid in a host methanotrophic bacterium may also be utilized. For example, an inducible/regulatable system of recombinant protein expression in methylotrophic and methanotrophic bacteria as described in, for example, U.S. Patent Appl. No. US 2010/0221813 may be used.

Methods of Producing a Desired Carbohydrate

[0062] The present disclosure provides a method of producing a carbohydrate by culturing a recombinant C₁ metabolizing microorganism of the present disclosure in the presence of methane (from any source), or a natural gas-derived carbon feedstock under conditions sufficient to produce the carbohydrate. In a specific embodiment, the present disclosure provides a method of producing a carbohydrate by culturing a recombinant C₁ metabolizing microorganism in the presence of a natural gas-derived carbon feedstock under conditions sufficient to produce the carbohydrate, wherein the C₁ metabolizing microorganism comprises an exogenous nucleic acid encoding a carbohydrate biosynthesis enzyme. Typically, the natural gas-derived carbon feedstock is natural gas, methane, or syngas. Conditions for culturing exemplary C₁ metabolizing microorganisms are illustrated in Example 1.

[0063] In a further embodiment, the present disclosure provides a method of producing a carbohydrate, said method comprising culturing a recombinant C₁ metabolizing microorganism in the presence of methane under conditions sufficient to produce the carbohydrate, wherein the C₁ metabolizing microorganism comprises an exogenous nucleic acid encoding a carbohydrate biosynthesis enzyme. In this embodiment, methane from any source is suitable for use in the practice of the present invention, including natural gas, bio-methane, and the like. As used herein, the term "bio-methane" refers to methane generated by fermentation of organic matter such as, for example, manure, waste water sludge, municipal solid waste, and the like, under anaerobic conditions.

[0064] A variety of culture methodologies may be used for the microorganisms described herein. For example, C₁ metabolizing microorganisms (such as methanotroph or methylotroph bacteria) may be grown by batch culture or continuous culture methodologies. In certain embodiments, the cultures are grown in a controlled culture unit, such as a fermentor, bioreactor, hollow fiber cell, or the like. Generally cells in log phase are often responsible for the bulk production of a product or intermediate of interest in some systems, whereas stationary or post-exponential phase production can be obtained in other systems.

[0065] A classical batch culturing method is a closed system in which the media composition is set when the culture is started and is not altered during the culture process. That is, media is inoculated at the beginning of the culturing process with one or more microorganisms of choice and then are allowed to grow without adding anything to the system. As used herein, a “batch” culture is in reference to not changing the amount of a particular carbon source initially added, whereas control of factors such as pH and oxygen concentration can be monitored and altered during the culture. In batch systems, metabolite and biomass compositions of the system change constantly up to the time the culture is terminated. Within batch cultures, cells (e.g., bacteria such as methylotrophs) will generally move from a static lag phase to a high growth logarithmic phase to a stationary phase where growth rate is reduced or stopped (and will eventually lead to cell death if conditions do not change).

[0066] A fed-batch system is a variation on the standard batch system in which a carbon substrate of interest is added in increments as the culture progresses. Fed-batch systems are useful when cell metabolism is likely to be inhibited by catabolite repression and when it is desirable to have limited amounts of substrate in the media. Since it is difficult to measure actual substrate concentration in fed-batch systems, an estimate is made based on changes of measureable factors such as pH, dissolved oxygen, and the partial pressure of waste gases. Batch and fed-batch culturing methods are common and known in the art (see, e.g., Thomas D. Brock, *Biotechnology: A Textbook of Industrial Microbiology*, 2nd Ed. (1989) Sinauer Associates, Inc., Sunderland, Mass.; Deshpande, *Appl. Biochem. Biotechnol.* 36:227, 1992).

[0067] Continuous cultures are “open” systems in the sense that defined culture media is continuously added to a bioreactor while an equal amount of used (“conditioned”) media is removed simultaneously for processing. Continuous cultures generally maintain the cells at a constant high, liquid phase density where cells are primarily in logarithmic growth phase. Alternatively, continuous culture may be practiced with immobilized cells (e.g., biofilm) where carbon and nutrients are continuously added and valuable products, by-products, and waste products are continuously removed from the cell mass. Cell immobilization may be achieved with a wide range of solid supports composed of natural materials, synthetic materials, or a combination thereof.

[0068] Continuous or semi-continuous culture allows for the modulation of one or more factors that affect cell growth or end product concentration. For example, one method may maintain a limited nutrient at a fixed rate (e.g., carbon source, nitrogen) and allow all other parameters to change over time. In other embodiments, several factors affecting growth may be continuously altered while cell concentration, as measured by media turbidity, is kept constant. The goal of a continuous culture system is to maintain steady state growth conditions while balancing cell loss due to media being drawn off against the cell growth rate. Methods of modulating nutrients and growth factors for continuous culture processes and techniques for maximizing the rate of product formation are well known in the art (see Brock, 1992).

[0069] Liquid phase bioreactors (e.g., stirred tank, packed bed, one liquid phase, two liquid phase, hollow fiber mem-

brane) are well known in the art and may be used for growth of non-naturally occurring microorganisms and biocatalysis.

[0070] By using gas phase bioreactors, substrates for bioproduction are absorbed from a gas by non-naturally occurring microorganisms, cell lysates or cell-free fractions thereof, rather than from a liquid. Use of gas phase bioreactors with microorganisms is known in the art (e.g., U.S. Pat. Nos. 2,793,096; 4,999,302; 5,585,266; 5,079,168; and 6,143,556; U.S. Statutory Invention Registration H1430; U.S. Patent Application Publication No. 2003/0032170; *Emerging Technologies in Hazardous Waste Management III*, 1993, eds. Tedder and Pohland, pp 411-428). Exemplary gas phase bioreactors include single pass system, closed loop pumping system, and fluidized bed reactor. By utilizing gas phase bioreactors, methane or other gaseous substrates are readily available for bioconversion by polypeptides with, for example, monooxygenase activity. In certain embodiments, methods for converting a gas into a carbohydrate are performed in gas phase bioreactors. In further embodiments, methods for converting a gas into a carbohydrate are performed in fluidized bed reactors. In a fluidized bed reactor, a fluid (i.e., gas or liquid) is passed upward through particle bed carriers, usually sand, granular-activated carbon, or diatomaceous earth, on which microorganisms can attach and grow. The fluid velocity is such that particle bed carriers and attached microorganisms are suspended (i.e., bed fluidization). The microorganisms attached to the particle bed carriers freely circulate in the fluid, allowing for effective mass transfer of substrates in the fluid to the microorganisms and increased microbial growth. Exemplary fluidized bed reactors include plug-flow reactors and completely mixed reactors. Uses of fluidized bed reactors with microbial biofilms are known in the art (e.g., Pfluger et al., *Bioresource Technol.* 102:9919, 2011; Fennell et al., *Biotechnol. Bioengin.* 40:1218, 1992; Ruggeri et al., *Water Sci. Technol.* 29:347, 1994; U.S. Pat. Nos. 4,032,407; 4,009,098; 4,009,105; and 3,846,289).

[0071] Recombinant C_1 metabolizing microorganisms described in the present disclosure may be grown as an isolated pure culture, with a heterologous non- C_1 metabolizing microorganism(s) that may aid with growth, or with one or more different strains or species of C_1 metabolizing microorganisms may be combined to generate a mixed culture.

[0072] In certain embodiments, carbohydrates of the present disclosure are produced during a specific phase of cell growth (e.g., lag phase, log phase, stationary phase, or death phase). It may be desirable for carbon from feedstock to be converted to the carbohydrate rather than to growth and maintenance of C_1 metabolizing microorganism. In some embodiments, non-naturally occurring C_1 metabolizing microorganism (e.g., methanotrophs, methylotrophs) as provided herein are cultured to a low to medium cell density (OD_{600}) and then production of carbohydrate is initiated. In some embodiments, a carbohydrate is produced while methanotrophic bacteria are no longer dividing or dividing very slowly. In some embodiments, the carbohydrate is produced only during stationary phase. In some embodiments, the carbohydrate is produced during log phase and stationary phase.

[0073] The fermenter composition comprising the carbohydrate produced by a recombinant C_1 metabolizing microorganism (e.g., methanotrophs, methylotrophs) provided herein may further comprise other organic compounds asso-

ciated with biological fermentation processes. For example, biological by-products of fermentation may include one or more of alcohols, epoxides, aldehydes, ketones, esters, or a combination thereof. In certain embodiments, the fermenter composition may contain one or more of the following alcohols: methanol, ethanol, butanol, or propanol. Other compounds, such as H₂O, CO, CO₂, CO N₂, H₂, O₂, and unutilized carbon feedstocks, such as methane, ethane, propane, and butane, may also be present in the fermenter off-gas.

[0074] In certain embodiments, the recombinant C₁ metabolizing microorganisms (e.g., methanotrophs, methylotrophs) provided herein produce a carbohydrate of the present invention at about 0.001 g/L of culture to about 500 g/L of culture. In some embodiments, the amount of carbohydrate produced is about 1 g/L of culture to about 100 g/L of culture. In some embodiments, the amount of carbohydrate produced is about 0.001 g/L, 0.01 g/L, 0.025 g/L, 0.05 g/L, 0.1 g/L, 0.15 g/L, 0.2 g/L, 0.25 g/L, 0.3 g/L, 0.4 g/L, 0.5 g/L, 0.6 g/L, 0.7 g/L, 0.8 g/L, 0.9 g/L, 1 g/L, 2.5 g/L, 5 g/L, 7.5 g/L, 10 g/L, 12.5 g/L, 15 g/L, 20 g/L, 25 g/L, 30 g/L, 35 g/L, 40 g/L, 45 g/L, 50 g/L, 60 g/L, 70 g/L, 80 g/L, 90 g/L, 100 g/L, 125 g/L, 150 g/L, 175 g/L, 200 g/L, 225 g/L, 250 g/L, 275 g/L, 300 g/L, 325 g/L, 350 g/L, 375 g/L, 400 g/L, 425 g/L, 450 g/L, 475 g/L, or 500 g/L.

Products

[0075] The present disclosure provides other useful products in addition to the recombinant C₁ metabolizing cells described herein. In one embodiment, the present disclosure provides a biomass comprising a recombinant C₁ metabolizing microorganism as described herein. In a specific embodiment, the present disclosure provides a biomass comprising a recombinant C₁ metabolizing microorganism, wherein the recombinant C₁ metabolizing microorganism comprises an exogenous nucleic acid encoding a carbohydrate biosynthesis enzyme and wherein the recombinant C₁ metabolizing microorganism is capable of converting a natural gas-derived feedstock into a desired carbohydrate. In a specific embodiment, the exogenous nucleic acid encodes a β -glucan biosynthesis enzyme, for example, a β -(1,3)-glucan synthase. In some embodiments, the biomass comprises a recombinant C₁ metabolizing microorganism and a desired carbohydrate, wherein the desired carbohydrate is a β -glucan and the recombinant C₁ metabolizing microorganism comprises an exogenous nucleic acid encoding a β -glucan biosynthesis enzyme, and wherein the C₁ metabolizing microorganism is capable of converting a natural gas-derived feedstock into a β -glucan. Exemplary β -glucans include a β -(1,3)-glucan, a β -(1,3)(1,6)-glucan, a β -(1,3)(1,4)-glucan, and a β -(1,4)-glucan. In certain embodiments, the desired carbohydrate is selected from a β -(1,3)-glucan, a β -(1,3)(1,6)-glucan, or a β -(1,3)(1,4)-glucan. In other embodiments, the desired carbohydrate is a β -(1,3)-glucan.

[0076] In a further embodiment, the present disclosure provides a biomass comprising a recombinant C₁ metabolizing microorganism, wherein the recombinant C₁ metabolizing microorganism comprises an exogenous nucleic acid encoding a carbohydrate biosynthesis enzyme and wherein the recombinant C₁ metabolizing microorganism is capable of converting methane into a desired carbohydrate. In a specific embodiment, the exogenous nucleic acid encodes a β -glucan biosynthesis enzyme, for example, a β -(1,3)-glucan synthase, and the C₁ metabolizing microorganism is

capable of converting methane into a β -glucan. Typically the β -glucan is selected from the group consisting of a β -glucan, such as, for example, a β -(1,3)-glucan, a β -(1,3)(1,6)-glucan, a β -(1,3)(1,4)-glucan, and a β -(1,4)-glucan. In certain embodiments, the desired carbohydrate is selected from the group consisting of a β -(1,3)-glucan, a β -(1,3)(1,6)-glucan, a β -(1,3)(1,4)-glucan. In other embodiments, the desired carbohydrate is a β -(1,3)-glucan.

[0077] As used herein, “biomass” refers to organic material having a biological origin, which may include one or more of whole cells, lysed cells, extracellular material, or the like. For example, the material harvested from a cultured microorganism (e.g., bacterial or yeast culture) is considered the biomass, which can include cells, cell membranes, cell cytoplasm, inclusion bodies, products secreted or excreted into the culture medium, or any combination thereof. In certain embodiments, biomass comprises the C₁ metabolizing microorganisms of this disclosure together with the media of the culture in which the C₁ metabolizing microorganisms of this disclosure were grown. In other embodiments, biomass comprises a C₁ metabolizing microorganism (whole or lysed or both) of this disclosure recovered from a culture grown on a C₁ substrate (e.g., natural gas, methane, and the like). In still other embodiments, biomass comprises the spent media supernatant from a culture of C₁ metabolizing microorganism cultured on a C₁ substrate. Such a culture may be considered a renewable resource. Biomass of the present invention is enriched with respect to levels of the desired carbohydrate.

[0078] Recombinant C₁ metabolizing microorganism of the present disclosure that are provided with a natural gas-derived substrate for cell growth are distinctive with respect to their carbon fingerprint as represented by their $\delta^{13}\text{C}$ values (as are the products derived from such recombinant C₁ metabolizing microorganisms). By way of background, stable isotopic measurements and mass balance approaches are widely used to evaluate global sources and sinks of methane (see Whiticar and Faber, *Org. Geochem.* 10:759, 1986; Whiticar, *Org. Geochem.* 16: 531, 1990). To use $\delta^{13}\text{C}$ values of residual methane to determine the amount oxidized, it is necessary to know the degree of isotopic fractionation caused by microbial oxidation of methane. For example, aerobic methanotrophs can metabolize methane through a specific enzyme, methane monooxygenase (MMO). Methanotrophs convert methane to methanol and subsequently formaldehyde. Formaldehyde can be further oxidized to CO₂ to provide energy to the cell in the form of reducing equivalents (NADH), or incorporated into biomass through either the RuMP or Serine cycles (Hanson and Hanson, *Microbiol. Rev.* 60:439, 1996), which are directly analogous to carbon assimilation pathways in photosynthetic organisms. More specifically, a Type I methanotroph uses the RuMP pathway for biomass synthesis and generates biomass entirely from CH₄, whereas a Type II methanotroph uses the serine pathway that assimilates 50-70% of the cell carbon from CH₄ and 30-50% from CO₂ (Hanson and Hanson, 1996). Methods for measuring carbon isotope compositions are provided in, for example, Templeton et al. (*Geochim. Cosmochim. Acta* 70:1739, 2006), which methods are hereby incorporated by reference in their entirety. Examples 2 describes the characterization of stable carbon isotope distribution in the cells of different C₁ metabolizing microorganisms. The highly negative $\delta^{13}\text{C}$ values for the cells was similarly reflected in the $\delta^{13}\text{C}$ of

compounds extracted from these cells, i.e., lipid fractions. The $\delta^{13}\text{C}$ of the invention products described herein (i.e., a recombinant C_1 metabolizing microorganism of the present disclosure as described herein), related biomass and carbohydrate compositions derived therefrom) can vary depending on the source and purity of the C_1 substrate used as demonstrated in Example 2.

[0079] In certain embodiments, a recombinant C_1 metabolizing microorganism of the present disclosure, and related biomass and carbohydrate compositions derived therefrom, exhibit a $\delta^{13}\text{C}$ of less than -30% , less than -31% , less than -32% , less than -33% , less than -34% , less than -35% , less than -36% , less than -37% , less than -38% , less than -39% , less than -40% , less than -41% , less than -42% , less than -43% , less than -44% , less than -45% , less than -46% , less than -47% , less than -48% , less than -49% , less than -50% , less than -51% , less than -52% , less than -53% , less than -54% , less than -55% , less than -56% , less than -57% , less than -58% , less than -59% , less than -60% , less than -61% , less than -62% , less than -63% , less than -64% , less than -65% , less than -66% , less than -67% , less than -68% , less than -69% , or less than -70% .

[0080] In certain embodiments, a recombinant C_1 metabolizing microorganism of the present disclosure, and related biomass and carbohydrate compositions derived therefrom, exhibit a $\delta^{13}\text{C}$ of about -35% to about -50% , -45% to about -35% , or about -50% to about -40% , or about -45% to about -65% , or about -60% to about -70% , or about -30% to about -70% .

[0081] In further embodiments, a C_1 metabolizing non-photosynthetic microorganism biomass has a $\delta^{13}\text{C}$ of less than about -30% , or ranges from about -40% to about -60% . In certain embodiments, the biomass comprises a recombinant C_1 metabolizing non-photosynthetic microorganism together with the spent media, or the biomass comprises a spent media supernatant composition from a culture of a recombinant C_1 metabolizing non-photosynthetic microorganism, wherein the $\delta^{13}\text{C}$ of the biomass is less than about -30% . In certain other embodiments, the carbohydrate composition is extracted or concentrated from a biomass, which can comprise recombinant C_1 metabolizing non-photosynthetic microorganisms together with the spent media from a culture, or a spent media supernatant composition from a culture of a recombinant C_1 metabolizing non-photosynthetic microorganism.

[0082] In certain embodiments, a carbohydrate composition derived from a C_1 metabolizing microorganism (which may optionally be an extract or isolate from the C_1 metabolizing microorganism biomass) comprises hydrogen, oxygen, and carbon atoms of at least about 50% to about 80% of the weight of the composition, and wherein the $\delta^{13}\text{C}$ of the composition is less than about -35% or less than about -36% or less than about -37% or less than about -38% or less than about -39% or less than about -40% . In certain embodiments, a carbohydrate composition derived therefrom comprises molecules having hydrogen, oxygen, and carbon atoms, wherein the hydrogen, oxygen, and carbon atoms are at least 50% , at least 55% , at least 60% , at least 65% , at least 70% , at least 75% , or at least 80% , or at least 90% , or at least 95% of the weight of the composition and wherein the $\delta^{13}\text{C}$ of the composition ranges from about -30% to about -70% , or wherein the $\delta^{13}\text{C}$ in the biomass decreases as cell density increases by about -5% to about -20% , or wherein the $\delta^{13}\text{C}$ of the biomass is higher than that

of CO_2 produced at the same time by an average of 5% to 15% when cultured in the presence or absence of copper.

[0083] Typically, a carbohydrate composition comprises a polysaccharide, and in some instances, it comprises a monosaccharide. In other embodiments the carbohydrate composition comprises a disaccharide. In some embodiments, the carbohydrate comprises a β -glucan. Typically, the β -glucan is a β -(1,3)-glucan. In other embodiments, the β -glucan is a β -(1,3)(1,6)-glucan, or a β -(1,3)(1,4)-glucan, or a β -(1,6)-glucan. Carbohydrate compositions derived from recombinant C_1 metabolizing microorganisms cultivated in the presence of a natural gas-derived substrate exhibit the $\delta^{13}\text{C}$ values described hereinabove.

[0084] Characterization of $\delta^{13}\text{C}$ of some C_1 metabolizing microorganisms cultivated in the presence of a natural gas-derived feedstock is illustrated in the examples, hereinbelow.

[0085] The present disclosure further provides an animal feed comprising the recombinant C_1 metabolizing microorganism, related biomass, and/or carbohydrate composition of the present disclosure. As contemplated in the practice of the present invention, the animal feed may be a livestock feed (such as, for example, pig feed, cattle feed, sheep feed, and the like), a poultry feed (such as, for example, chicken feed, turkey feed, and the like), or a fish feed (such as, for example, salmon feed, shell fish feed, and the like). The animal feed may further comprise an additive, such as, for example, a plant-derived material (including, for example, those derived from grains such as, for example, corn, barley, oats, rice, rye, wheat, sorghum, Brewer's spent grain, and the like; and those derived from legumes, such as, for example, alfalfa, clover, peas, beans, lentils, soybeans, and the like), an animal-derived material (such as, for example, fish meal), and/or a microorganism-derived material (including, for example, biomass from a heterologous microorganism that may be, for example, a bacteria, a yeast, or an algae). In some embodiments, the plant-derived material additive is soy meal or pea protein.

[0086] In a further embodiment, the present disclosure provides a culture or fermentation medium comprising the recombinant C_1 metabolizing microorganism, related biomass, and/or carbohydrate composition of the present disclosure. Typically, the culture or fermentation medium further comprises an amino acid and/or water. In an additional embodiment, the present disclosure provides a cell culture composition comprising a culture or fermentation medium as described herein, and a second microorganism. Typically, a second microorganism is a bacteria, a yeast, or an algae.

[0087] Embodiments of the present invention include the following:

[0088] 1. A biomass derived from a culture of a recombinant C_1 metabolizing microorganism, wherein the recombinant microorganism comprises an exogenous nucleic acid encoding a carbohydrate biosynthesis enzyme, wherein the recombinant C_1 metabolizing is capable of converting a natural gas-derived carbon feedstock into a desired carbohydrate.

[0089] 2. A biomass derived from a culture of a recombinant C_1 metabolizing microorganism, wherein the recombinant C_1 metabolizing microorganism comprises an exogenous nucleic acid encoding a carbohydrate biosynthesis enzyme, wherein the recombinant C_1 metabolizing microorganism is capable of converting methane into a desired carbohydrate.

- [0090] 3. The biomass of any of embodiments 1-2, wherein the recombinant C_1 metabolizing microorganism is a non-photosynthetic C_1 metabolizing microorganism.
- [0091] 4. The biomass of any of embodiments 1-3, wherein the carbohydrate is selected from the group consisting of a polysaccharide, a disaccharide, and a monosaccharide.
- [0092] 5. The biomass of embodiment 4, wherein the carbohydrate is a monosaccharide.
- [0093] 6. The biomass of embodiment 4, wherein the carbohydrate is a disaccharide.
- [0094] 7. The biomass of embodiment 4, wherein the carbohydrate is a polysaccharide.
- [0095] 8. The biomass of embodiment 7, wherein the polysaccharide is a β -glucan.
- [0096] 9. The biomass of embodiment 8, wherein the β -glucan is β -(1,3)-glucan.
- [0097] 10. The biomass of embodiment 8, wherein the β -glucan is β -(1,3)(1,6)-glucan.
- [0098] 11. The biomass of embodiment 8, wherein the β -glucan is β -(1,3)(1,4)-glucan.
- [0099] 12. The biomass of embodiment 8, wherein the β -glucan is β -(1,4)-glucan.
- [0100] 13. The biomass of embodiment 8, wherein the β -glucan is β -(1,6)-glucan.
- [0101] 14. The biomass of any of embodiments 1-13, wherein the sequence of the exogenous nucleic acid is codon optimized for optimal expression from the recombinant C_1 metabolizing microorganism.
- [0102] 15. The biomass of any of embodiments 1-14, wherein the exogenous nucleic acid encodes a gluconeogenesis enzyme.
- [0103] 16. The biomass of embodiment 15, wherein the gluconeogenesis enzyme is selected from the group consisting of a pyruvate carboxylase, a phosphoenolpyruvate carboxykinase, an enolase, a phosphoglycerate mutase, a phosphoglycerate kinase, a glyceraldehyde-3-phosphate dehydrogenase, a Type A aldolase, a fructose 1,6-bisphosphatase, a phosphofructokinase, a phosphoglucose isomerase, a hexokinase, and a glucose-6-phosphate.
- [0104] 17. The biomass of any of embodiments 1-14, wherein the exogenous nucleic acid encodes a glycogenesis enzyme.
- [0105] 18. The biomass of embodiment 17, wherein the glycogenesis enzyme is selected from the group consisting of a glucose-1-phosphate adenylyltransferase, a glycogen synthase, and a 1,4-alpha-glucan-branching protein.
- [0106] 19. The biomass of any of embodiments 8-14, wherein the exogenous nucleic acid is a β -glucan synthase.
- [0107] 20. The biomass of any of embodiments 1-19, wherein the exogenous nucleic acid encodes a carbohydrate biosynthesis enzyme that is endogenous to a bacteria.
- [0108] 21. The biomass of any of embodiments 1-19, wherein the exogenous nucleic acid encodes a carbohydrate biosynthesis enzyme that is endogenous to an organism selected from the group consisting of a yeast, a fungi, and a plant.
- [0109] 22. The biomass of any of embodiments 1-19, wherein the exogenous nucleic acid encodes a carbohydrate biosynthesis enzyme that is endogenous to a microorganism selected from the group consisting of *E. coli* and *C. glutamicum*.
- [0110] 23. The biomass of any of embodiments 1-14, wherein the exogenous nucleic acid encodes a carbohydrate biosynthesis enzyme selected from the group consisting of any of SEQ ID NOs:2, 4, 6, 8 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, and 38.
- [0111] 24. The biomass of any of embodiments 1-23, wherein the exogenous nucleic acid encoding carbohydrate biosynthesis pathway enzyme is operatively linked to an expression control sequence.
- [0112] 25. The biomass of embodiment 24, wherein the expression control sequence is an exogenous expression control sequence.
- [0113] 26. The biomass of any of embodiments 1-25, wherein the C_1 metabolizing microorganism further comprises a deletion of an endogenous enzyme activity. 27. The biomass according to any of embodiments 1-26, wherein the C_1 metabolizing microorganism is a methanotroph.
- [0114] 28. The biomass according to embodiment 27, wherein the methanotroph is *Methylomonas*, *Methylobacter*, *Methylococcus*, *Methylosinus*, *Methylocystis*, *Methylomicrobium*, *Methanomonas*, *Methylocella*, or *Methylocapsa*.
- [0115] 29. The biomass of embodiment 27, wherein the methanotroph is selected from the group consisting of *Methylococcus capsulatus* Bath strain, *Methylomonas methanica* 16a (ATCC PTA 2402), *Methylosinus trichosporium* OB3b (NRRL B-11,196), *Methylosinus sporium* (NRRL B-11,197), *Methylocystis parvus* (NRRL B-11,198), *Methylomonas methanica* (NRRL B-11,199), *Methylomonas albus* (NRRL B-11,200), *Methylobacter capsulatus* (NRRL B-11,201), *Methylobacterium organophilum* (ATCC 27,886), *Methylomonas* sp AJ-3670 (FERM P-2400), *Methylocella silvestris*, *Methylocella palustris* (ATCC 700799), *Methylocella tundrae*, *Methylocystis daltona* strain SB2, *Methylocystis bryophila*, *Methylocapsa aurea* KYG, *Methylacidiphilum inferorum*, *Methylibium petroleiphilum*, and *Methylomicrobium alcahphilum*.
- [0116] 30. The biomass according to any one of embodiments 1 and 3-29, wherein the natural gas-derived carbon feedstock is selected from the group consisting of natural gas, syngas, methane, methanol, formaldehyde, formic acid, carbon monoxide, carbon dioxide, cyanide, a methylamine, a methylthiol, a methylhalogen, and any combination or two or more thereof.
- [0117] 31. The biomass of embodiment 30, wherein the natural gas-derived carbon feedstock is natural gas.
- [0118] 32. The biomass of any of embodiments 1, and 3-30, wherein the natural gas-derived carbon feedstock is methane.
- [0119] 33. The biomass of embodiment 30, wherein the natural gas-derived carbon feedstock is syngas.
- [0120] 34. The biomass of embodiment 30, wherein the C_1 metabolizing microorganism is a syngas metabolizing bacteria.
- [0121] 35. The biomass according to embodiment 34, wherein the syngas metabolizing bacteria is selected from the group consisting of *Clostridium autoethanogenum*, *Clostridium ljungdahli*, *Clostridium ragsdalei*, *Clostridium carboxydivorans*, *Butyridbacterium methylophilum*, *Clostridium woodii*, and *Clostridium neopropanologen*.
- [0122] 36. The biomass according to any one of embodiments 1 and 3-35, wherein the $\delta^{13}\text{C}$ of the biomass is less than -40% .
- [0123] 37. The biomass of embodiment 2, wherein the methane is bio-methane.

- [0124] 38. A composition comprising a carbohydrate composition, wherein the carbohydrate composition exhibits a $\delta^{13}\text{C}$ of less than -40% .
- [0125] 39. The composition of embodiment 38, wherein the carbohydrate comprises a β -glucan.
- [0126] 40. The composition of embodiment 39, wherein the β -glucan is β -(1,3)-glucan.
- [0127] 41. An animal feed comprising the biomass of any of embodiments 1-37 or the composition of any of embodiments 38-40.
- [0128] 42. The animal feed of embodiment 41, further comprising a plant-derived material.
- [0129] 43. The animal feed of embodiment 41, wherein the plant-derived material is selected from the group consisting of soybean meal and pea protein.
- [0130] 44. A culture or fermentation medium comprising the biomass of any of embodiments 1-37 or the composition of any of embodiments 38-40.
- [0131] 45. A recombinant C_1 metabolizing microorganism, wherein the recombinant microorganism comprises an exogenous nucleic acid encoding a carbohydrate biosynthesis enzyme, wherein the recombinant C_1 metabolizing microorganism is capable of converting a natural gas-derived carbon feedstock into a desired carbohydrate.
- [0132] 46. A recombinant C_1 metabolizing microorganism, wherein the recombinant C_1 metabolizing microorganism comprises an exogenous nucleic acid encoding a carbohydrate biosynthesis enzyme, wherein the recombinant C_1 metabolizing microorganism is capable of converting methane into a desired carbohydrate.
- [0133] 47. The recombinant C_1 metabolizing microorganism of any of embodiments 45-46, wherein the recombinant C_1 metabolizing microorganism is a non-photosynthetic C_1 metabolizing microorganism.
- [0134] 48. The recombinant C_1 metabolizing microorganism of any of embodiments 45-47, wherein the carbohydrate is selected from the group consisting of a polysaccharide, a disaccharide, and a monosaccharide.
- [0135] 49. The recombinant C_1 metabolizing microorganism of embodiment 48, wherein the carbohydrate is a monosaccharide.
- [0136] 50. The recombinant C_1 metabolizing microorganism of embodiment 48, wherein the carbohydrate is a disaccharide.
- [0137] 51. The recombinant C_1 metabolizing microorganism of embodiment 48, wherein the carbohydrate is a polysaccharide.
- [0138] 52. The recombinant C_1 metabolizing microorganism of embodiment 51, wherein the polysaccharide is a β -glucan.
- [0139] 53. The recombinant C_1 metabolizing microorganism of embodiment 52, wherein the β -glucan is β -(1,3)-glucan.
- [0140] 54. The recombinant C_1 metabolizing microorganism of embodiment 52, wherein the β -glucan is β -(1,3)(1,6)-glucan.
- [0141] 55. The recombinant C_1 metabolizing microorganism of embodiment 52, wherein the β -glucan is β -(1,3)(1,4)-glucan.
- [0142] 56. The recombinant C_1 metabolizing microorganism of embodiment 52, wherein the β -glucan is β -(1,4)-glucan.
- [0143] 57. The recombinant C_1 metabolizing microorganism of embodiment 52, wherein the β -glucan is β -(1,6)-glucan.
- [0144] 58. The recombinant C_1 metabolizing microorganism of any of embodiments 45-57, wherein the sequence of the exogenous nucleic acid is codon optimized for optimal expression from the recombinant C_1 metabolizing microorganism.
- [0145] 59. The recombinant C_1 metabolizing microorganism of any of embodiments 45-58, wherein the exogenous nucleic acid encodes a gluconeogenesis enzyme.
- [0146] 60. The recombinant C_1 metabolizing microorganism of embodiment 59, wherein the gluconeogenesis enzyme is selected from the group consisting of a pyruvate carboxylase, a phosphoenolpyruvate carboxykinase, an enolase, a phosphoglycerate mutase, a phosphoglycerate kinase, a glyceraldehyde-3-phosphate dehydrogenase, a Type A aldolase, a fructose 1,6-bisphosphatase, a phosphofructokinase, a phosphoglucose isomerase, a hexokinase, and a glucose-6-phosphate.
- [0147] 61. The recombinant C_1 metabolizing microorganism of any of embodiments 45-58, wherein the exogenous nucleic acid encodes a glycogenesis enzyme.
- [0148] 62. The recombinant C_1 metabolizing microorganism of embodiment 61, wherein the glycogenesis enzyme is selected from the group consisting of a glucose-1-phosphate adenylyltransferase, a glycogen synthase, and a 1,4-alpha-glucan-branching protein.
- [0149] 63. The recombinant C_1 metabolizing microorganism of any of embodiments 52-57, wherein the exogenous nucleic acid is a β -glucan synthase.
- [0150] 64. The recombinant C_1 metabolizing microorganism of any of embodiments 45-63, wherein the exogenous nucleic acid encodes a carbohydrate biosynthesis enzyme that is endogenous to a bacteria.
- [0151] 65. The recombinant C_1 metabolizing microorganism of any of embodiments 45-63, wherein the exogenous nucleic acid encodes a carbohydrate biosynthesis enzyme that is endogenous to an organism selected from the group consisting of a yeast, a fungi, and a plant.
- [0152] 66. The recombinant C_1 metabolizing microorganism of any of embodiments 45-63, wherein the exogenous nucleic acid encodes a carbohydrate biosynthesis enzyme that is endogenous to a microorganism selected from the group consisting of *E. coli*, and *C. glutamicum*.
- [0153] 67. The recombinant C_1 metabolizing microorganism of any of embodiments 45-57, wherein the exogenous nucleic acid encodes a carbohydrate biosynthesis enzyme selected from the group consisting of any of SEQ ID NOs: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, and 38.
- [0154] 68. The recombinant C_1 metabolizing microorganism of any of embodiments 45-67, wherein the exogenous nucleic acid encoding carbohydrate biosynthesis pathway enzyme is operatively linked to an expression control sequence.
- [0155] 69. The recombinant C_1 metabolizing microorganism of embodiment 68, wherein the expression control sequence is an exogenous expression control sequence.
- [0156] 70. The recombinant C_1 metabolizing microorganism of any of embodiments 45-69, wherein the C_1 metabolizing microorganism further comprises a deletion of an endogenous enzyme activity.

[0157] 71. The recombinant C₁ metabolizing microorganism according to any of embodiments 45-70, wherein the C₁ metabolizing microorganism is a methanotroph.

[0158] 72. The recombinant C₁ metabolizing microorganism according to embodiment 71, wherein the methanotroph is *Methylomonas*, *Methylobacter*, *Methylococcus*, *Methylosinus*, *Methylocystis*, *Methylomicrobium*, *Methanomonas*, *Methylocella*, or *Methylocapsa*.

[0159] 73. The recombinant C₁ metabolizing microorganism of embodiment 71, wherein the methanotroph is selected from the group consisting of *Methylococcus capsulatus* Bath strain, *Methylomonas methanica* 16a (ATCC PTA 2402), *Methylosinus trichosporium* OB3b (NRRL B-11,196), *Methylosinus sporium* (NRRL B-11,197), *Methylocystis parvus* (NRRL B-11,198), *Methylomonas methanica* (NRRL B-11,199), *Methylomonas albus* (NRRL B-11,200), *Methylobacter capsulatus* (NRRL B-11,201), *Methylobacterium organophilum* (ATCC 27,886), *Methylomonas* sp AJ-3670 (FERM P-2400), *Methylocella silvestris*, *Methylocella palustris* (ATCC 700799), *Methylocella tundrae*, *Methylocystis daltona* strain SB2, *Methylocystis bryophila*, *Methylocapsa aurea* KYG, *Methylacidiphilum infernorum*, *Methylibium petroleiphilum*, and *Methylomicrobium alcaliphilum*.

[0160] 74. The recombinant C₁ metabolizing microorganism according to any one of embodiments 45 and 47-73 wherein the natural gas-derived carbon feedstock is selected from the group consisting of natural gas, syngas, methane, methanol, formaldehyde, formic acid, carbon monoxide, carbon dioxide, cyanide, a methylamine, a methylthiol, a methylhalogen, and any combination or two or more thereof.

[0161] 75. The recombinant C₁ metabolizing microorganism of embodiment 74, wherein the natural gas-derived carbon feedstock is natural gas.

[0162] 76. The recombinant C₁ metabolizing microorganism of embodiment 74, wherein the natural gas-derived carbon feedstock is methane.

[0163] 77. The recombinant C₁ metabolizing microorganism of embodiment 74, wherein the natural gas-derived carbon feedstock is syngas.

[0164] 78. The recombinant C₁ metabolizing microorganism of embodiment 77, wherein the C₁ metabolizing microorganism is a syngas metabolizing bacteria.

[0165] 79. The biomass according to embodiment 78, wherein the syngas metabolizing bacteria is selected from the group consisting of *Clostridium autoethanogenum*, *Clostridium ljungdahli*, *Clostridium ragsdalei*, *Clostridium carboxydvorans*, *Butyridbacterium methylotrophicum*, *Clostridium woodii*, and *Clostridium neopropanologen*.

[0166] 80. The recombinant C₁ metabolizing microorganism according to any one of embodiments 45 and 47-79, wherein the $\delta^{13}\text{C}$ of the biomass is less than -40%.

[0167] 81. The recombinant C₁ metabolizing microorganism of embodiment 46, wherein the methane is bio-methane.

[0168] 82. A method of producing a carbohydrate, said method comprising culturing the recombinant C₁ metabolizing microorganism of any of embodiments 45 and 47-68 in the presence of a natural gas-derived carbon feedstock under conditions sufficient to produce the carbohydrate.

[0169] 83. A method of producing a carbohydrate, said method comprising culturing the recombinant C₁ metabolizing microorganism of embodiment 46 in the presence of a methane under conditions sufficient to produce the carbohydrate.

[0170] 84. The method of embodiment 83, wherein the carbohydrate is a β -glucan.

[0171] 85. A carbohydrate produced by the method of embodiment 82, wherein the carbohydrate exhibits a $\delta^{13}\text{C}$ in the range of from about -40‰ to about -60‰.

[0172] The foregoing and other aspects of the invention may be better understood in connection with the following, non-limiting examples.

EXAMPLES

Example 1

Culture and Bioreactor Conditions for C₁ Metabolizing Microorganisms

[0173] Exemplary C₁ metabolizing microorganisms of the instant disclosure (methanotrophs, methylotrophs, clostridia) were cultured in tubes, in vials, in bottles, on plates, or in a bioreactor (fermentation). Growth conditions, media, and carbon source for various microorganisms are described in this example.

Methylosinus trichosporium Strain OB3b (NCIMB 11131); *Methylomonas* sp. Strain 16a (ATCC PTA-2402); or *Methylomonas methanica*

[0174] For serum bottles, the bacteria were cultured at 30° C. in Higgins minimal nitrate salts medium (NSM; Cornish et al., *J. Gen. Microbiol.* 130:2565, 1984; Park et al., *Biotechnol. Bioeng.* 38:423, 1991) or MM-W1 medium. The headspace composition was adjusted to a 1:1 volume of methane:air. The bottles were shaken at a rate of 200-250 rpm. Alternatively, the culture was maintained on NSM-media plates containing 1.5% w/v agar grown in a gas-tight chamber containing a 1:1 (v/v) methane:air gas mixture, or in the presence of methanol vapor (via 0.5 mL methanol in the lid of parafilm-sealed plates) or on NSM-media plates supplemented with 0.5% methanol. Plates were incubated inverted in a humidified chamber at 30° C.

[0175] The composition of the NSM medium used was as follows: 1.0 g MgSO₄*7H₂O, 0.20 g CaCl₂*6H₂O, 2.0 ml chelated iron solution (0.1 g ferric (III) ammonium citrate or 0.5 g ferric (III) chloride; 0.2 g EDTA, sodium salt; 0.3 ml HCl, concentrated; 100.0 ml distilled deionized H₂O), 1.0 g KNO₃, 0.5 ml trace element solution (500.0 mg EDTA, 200.0 mg FeSO₄. 7H₂O, 10.0 mg ZnSO₄*7H₂O, 3.0 mg MnCl₂*4H₂O, 30.0 mg H₃BO₃, 20.0 mg CoCl₂*6H₂O, 1.0 mg CaCl₂*2H₂O, 2.0 mg NiCl₂*6H₂O, 3.0 mg Na₂MoO₄*2H₂O, 1.0 L distilled water), 0.272 g KH₂PO₄, 0.717 g Na₂HPO₄*12H₂O, optionally 12.5 g purified agar (e.g., Oxoid L28 or Bacto™ agar; used when making plates), 1.0 L distilled deionized water, pH adjusted to 6.8 and autoclaved at 121° C. for 15 minutes.

[0176] For fermentation, a 2-liter bioreactor containing 1 L of sterilized defined media MM-W1 was inoculated with cells from serum bottle batch cultures (10-20% v/v) grown in MM-W1 supplied with a 1:1 (v/v) mixture of methane and air. The composition of medium MM-W1 used was as follows: 0.8 mM MgSO₄*7H₂O, 10 mM NaNO₃, 0.14 mM CaCl₂, 1.2 mM NaHCO₃, 2.35 mM KH₂PO₄, 3.4 mM K₂HPO₄, 20.7 μM Na₂MoO₄*2H₂O, 1 μM CuSO₄*5H₂O, 10 μM Fe^{III}-Na-EDTA, and 1 mL per liter of trace metals solution (containing, per liter 500 mg FeSO₄*7H₂O, 400 mg ZnSO₄*7H₂O, 20 mg MnCl₂*7H₂O, 50 mg CoCl₂*6H₂O, 10 mg NiCl₂*6H₂O, 15 mg H₃BO₃, 250 mg EDTA). Phosphate, bicarbonate, and Fe^{III}-Na-EDTA were added after the

media was autoclaved and cooled. Bicarbonate was added up to 0.1% (w/v) in certain fermentations. The reactor contents were stirred with an overhead impeller at a constant 750 rpm. The culture was fed with a constant methane sparging at about 60 mL/min to about 120 mL/min, while concentrated oxygen (at least 85%) was supplied at a variable rate of about 10-100 mL/min to maintain a dissolved oxygen level of about 40% to about 80% (relative to air saturation of the media).

[0177] Temperature in the bioreactor was maintained at 30° C. and pH was maintained at 7.1±0.1 using automated addition of 0.5M NaOH and 0.5M HCl, along with other additions, to the culture about every 4 hours to about 24 hours (corresponding to an OD₆₀₀ increase of approximately 5 OD units). The other additions alternated between a metal addition (10 µM CuSO₄, 5 µM FeSO₄, 5 µM Fe^{III}-Na-EDTA final concentrations) and a nutrient addition (5.75 mM K_xH_yPO₄, 10 mM NaNO₃). Under these conditions, essentially linear growth was observed, with an effective biomass generation rate of about 2.7 to about 3.3 grams dry cell weight per liter per day to an OD₆₀₀ of greater than 20. Culture biomass was harvested by centrifugation, washed once in MM-W 1 media, and recovered biomass was either frozen at -80° C. or used immediately for fractionation of cellular components (e.g., lipid extraction).

[0178] A semi-continuous fermentation approach can also be applied to maintain biomass productivity and reduce time associated with fermentation shut-down and start-up (i.e., turn-around time or lead time).

[0179] Harvesting of the bacterial biomass was performed at approximately 12-24 hour intervals, as the culture density approached (but before entering) stationary phase. Approximately half of the bioreactor volume was removed by transferring to a separate container via centrifugal pump. An equal volume of sterilized or recycled media was then returned to the bioreactor such that the optical density of the reactor was approximately half of its initial value. The bioreactor fermentation was continued according to the above protocol so that multiple cycles of growth and biomass recovery could be carried out during a single fermentation run.

Methylococcus capsulatus Bath (NCIMB 11132)

[0180] The bacteria were cultured at 42° C. in serum bottles containing Higgins minimal nitrate salts medium (NSM) or MM-W1 medium. The headspace composition was adjusted to a 1:1 volume of methane:air. The bottles were shaken at a rate of 200-250 rpm. Alternatively, the culture was maintained on NSM-media plates solidified with 1.5% w/v agar grown in a gas-tight chamber containing a 1:1 (v/v) methane:air gas mixture. Plates were incubated inverted in the chamber at 42° C.

[0181] For fermentation, a 3-liter bioreactor containing 1.25 L sterilized media MMF1.1 was inoculated with cells from serum bottle batch cultures (10-20% v/v) grown in the same media supplied with a 1:1 (v/v) mixture of methane and air. The composition of medium MMF1.1 was as follows: 0.8 mM MgSO₄*7H₂O, 40 mM NaNO₃, 0.14 mM CaCl₂, 6 mM NaHCO₃, 4.7 mM KH₂PO₄, 6.8 mM K₂HPO₄, 20.7 µM Na₂MoO₄*2H₂O, 6 µM CuSO₄*5H₂O, 10 µM Fe^{III}-Na-EDTA, and 1 mL per liter of trace metals solution (containing, per liter 500 mg FeSO₄*7H₂O, 400 mg ZnSO₄*7H₂O, 20 mg MnCl₂*7H₂O, 50 mg CoCl₂*6H₂O, 10 mg NiCl₂*6H₂O, 15 mg H₃BO₃, 250 mg EDTA). Phosphate, bicarbonate, and Fe^{III}-Na-EDTA were added after

media was autoclaved and cooled. The reactor contents were stirred with an overhead impeller at a constant 750 rpm. The culture was fed with a constant methane sparging at about 60 to about 200 mL/min, while concentrated oxygen (>85%) was supplied at a variable rate of 15-90 mL/min and the dissolved oxygen level was maintained below 10% (relative to air saturation of the media).

[0182] Temperature in the bioreactor was maintained at 44° C. and pH was maintained at 7.0±0.1 using automated addition of 0.5M NaOH and 0.5M HCl, along with additions of copper and iron (5 µM CuSO₄, 5 µM FeSO₄, 10 µM Fe^{III}-Na-EDTA final concentration) to the culture every 3-6 hours (corresponding to an OD₆₀₀ increase of approximately 3-5 OD units after reaching OD 5). Under these conditions, essentially linear growth was observed, with effective biomass generation rate of more than 5 grams dry cell weight per liter per day to an OD₆₀₀ of greater than 10. Culture biomass was harvested by centrifugation, the cells washed once in MM-W1 media and cell pellets were either frozen at -80° C. or used immediately for fractionation of cellular components.

[0183] Nutrient depletion was recognized as an issue that could limit the growth yield during fermentation. To avoid limitation of nutrients, mainly nitrogen and phosphate, nutrient feeds composed of 2-fold concentrated MMF1.1 were initiated after culture OD₆₀₀ exceeded 5. The nutrient feed was initiated at dilution rates corresponding to approximately half of the cultures' growth rate to avoid wash-out and to maintain an increase in OD while expanding the culture volume. The bioreactor fermentation was continued according to the above protocol so that multiple cycles of growth and biomass recovery could be carried out during a single fermentation run.

Methylobacterium extorquens or *Methylosinus trichosporium* Strain OB3b (NCIMB 11131)

[0184] The bacteria is cultured at 30° C. in tubes containing Higgins minimal nitrate salts medium (NSM) supplemented with 0.5% methanol. The tubes are shaken at a rate of 200-250 rpm. Alternatively, the cultures are maintained on NSM-media plates containing 1.5% w/v agar grown in the presence of methanol vapor (via 0.5 mL methanol in the lid of parafilm-sealed plates) or supplemented with 0.5% methanol. Plates are incubated inverted in a humidified chamber under normal atmosphere at 30° C.

[0185] For fermentation, a 2-liter bioreactor containing 1 L defined media MM-W1 is inoculated with cells from culture tube batch culture (10-20% v/v). The composition of medium MM-W1 was as described above. The reactor contents are stirred with an overhead impeller at a constant 800 rpm. The culture is fed with an initial bolus of methanol to a final concentration of 0.5% and variable methanol feed, while pure oxygen was supplied at a variable rate of 30-100 mL/min to maintain a dissolved oxygen level of 60-90% (relative to air saturation of the media).

[0186] Temperature in the bioreactor was maintained at 30° C. and pH was maintained at 7.1±0.1 using automated addition of 0.5M NaOH and 1M HCl, along with the metal and nutrient additions as described above. Under these conditions, essentially linear growth is observed, with effective biomass generation rate 2.7 to 3.3 grams dry cell weight per liter per day to an OD₆₀₀ of greater than 20. Culture biomass was harvested by centrifugation, the cells washed

once in MM-W 1 media and cell pellets were either frozen at -80°C . or used immediately for fractionation of cellular components.

[0187] A semi-continuous fermentation approach can also be applied to maintain biomass productivity and reduce time associated with fermentation shut-down and start-up (i.e., turn-around time or lead time).

[0188] Harvesting of the accumulated bacterial biomass was performed at approximately 12-24 hour intervals, as the culture density approached (but before entering) stationary phase. Approximately half of the bioreactor volume was removed by transferring to a separate container via centrifugal pump. An equal volume of fresh or recycled media was then returned to the bioreactor such that the optical density of the reactor was approximately half of its initial value. The bioreactor fermentation was continued according to the above protocol so that multiple cycles of growth and biomass recovery was carried out during a single fermentation run.

Clostridium autoethanogenum and *Clostridium ljungdahlii*

[0189] The *Clostridium* bacteria are cultivated anaerobically in 100 mL modified PETC medium (ATCC medium 1754) at 37°C . in plastic-coated 500 ml-Schott Duran® GL45 bottles with butyl rubber stoppers and 200 kPa steel mill waste gas. Growth is monitored by measuring the optical density at 600 nm (OD_{600}).

[0190] The modified PETC medium contains (per liter) 1 g NH_4Cl , 0.4 g KCl , 0.2 g $\text{MgSO}_4 \cdot 7\text{H}_2\text{O}$, 0.8 g NaCl , 0.1 g KH_2PO_4 , 20 mg $\text{CaCl}_2 \cdot 2\text{H}_2\text{O}$, 10 ml trace elements solution (see below), 10 ml Wolfe's vitamin solution (see below), 2 g NaHCO_3 , and 1 mg resazurin. After the pH is adjusted to 5.6, the medium is boiled, dispensed anaerobically, and autoclaved at 121°C . for 15 min. Steel mill waste gas (composition: 44% CO , 32% N_2 , 22% CO_2 , 2% H_2) or equivalent synthetic mixtures are used as a carbon source. The media has a final pH of 5.9 and is reduced with cysteine-HCl and Na_2S at a concentration of 0.008% (w/v).

[0191] The trace elements solution contains 2 g nitrilotriacetic acid (adjusted to pH 6 with KOH before addition of the remaining ingredients), 1 g MnSO_4 , 0.8 g $\text{Fe}(\text{SO}_4)_2 (\text{NH}_4)_2 \cdot 6\text{H}_2\text{O}$, 0.2 g $\text{CoCl}_2 \cdot 6\text{H}_2\text{O}$, 0.2 mg $\text{ZnSO}_4 \cdot 7\text{H}_2\text{O}$, 20 mg $\text{CuCl}_2 \cdot 2\text{H}_2\text{O}$, 20 mg $\text{NiCl}_2 \cdot 6\text{H}_2\text{O}$, 20 mg $\text{Na}_2\text{MoO}_4 \cdot 2\text{H}_2\text{O}$, 20 mg Na_2SeO_4 , and 20 mg Na_2WO_4 per liter.

[0192] Wolfe's vitamin solution (Wolin et al., *J. Biol. Chem.* 238:2882, 1963) contains (per liter) 2 mg biotin, 2 mg folic acid, 10 mg pyridoxine hydrochloride, 5 mg thiamine-HCl, 5 mg riboflavin, 5 mg nicotinic acid, 5 mg calcium D-(+)-pantothenate, 0.1 mg vitamin B12, 5 mg p-aminobenzoic acid, and 5 mg thioctic acid.

[0193] a. *Clostridium autoethanogenum* Fermentation

[0194] Fermentation of *Clostridium autoethanogenum* is conducted using methods similar to those described in, for example, U.S. Patent Appl. No. 2011/0300593. Briefly, a 2-liter bioreactor containing 1.3 L Solution A (3.083 g NH_4Ac ; 0.61 g $\text{MgCl}_2 \cdot 6\text{H}_2\text{O}$; 0.294 g $\text{CaCl}_2 \cdot 2\text{H}_2\text{O}$; 0.15 g KCl ; 0.12 g NaCl (optional); up to 1 L with distilled water) is sparged with N_2 gas. An 85% solution of H_3PO_4 (2.025 mL, 30 mM) is added and the pH adjusted to 5.3 using concentrated, aqueous NH_4OH . Then 13.5 mL Solution B (20.0 mg Biotin; 20.0 mg Folic acid; 10.0 mg pyridoxine HCl; 50.0 mg thiamine*HCl; 50.0 mg Riboflavin; 50.0 mg nicotinic acid; 50.0 mg calcium D-(*)-pantothenate; 50.0 mg vitamin B12; 50.0 mg p-aminobenzoic acid; 50.0 mg thio-

ctic acid; up to 1 L with distilled water) is added and the solution sparged with N_2 gas. Chromium (II) chloride is added until the oxidation-reduction potential (ORP) of the solution decreases to approximately -200 mV , wherein resazurin (1.35 mL of a 2 g/L solution) is added. Sodium polysulfide (5.4 mL of a 3M solution, see below) is added and the solution sparged with N_2 and then CO containing gas (1% H_2 ; 13% N_2 ; 71% CO ; 15% CO_2). A metal sulfide solution (150 mL, see below) is added and the solution sparged a further 30 minutes, before inoculation with an actively growing *C. autoethanogenum* culture at a level of approximately 5% (v/v).

[0195] The sodium polysulfide solution is prepared in a 500 ml flask that is charged with Na_2S (93.7 g, 0.39 mol) and 200 ml H_2O . The solution is stirred until the salt dissolves and sulfur (25 g, 0.1 mol) is added under constant N_2 flow. After stirring at room temperature for 2 hours, the sodium polysulfide solution (about 4 M with respect to Na and about 5 M with respect to sulfur), now a clear reddish brown liquid, is transferred into N_2 purged serum bottles, and wrapped in aluminum foil.

[0196] The chromium (II) solution is prepared in a 1 L three necked flask that is fitted with a gas tight inlet and outlet to allow working under inert gas and subsequent transfer of the desired product into a suitable storage flask. The flask is charged with $\text{CrCl}_3 \cdot 6\text{H}_2\text{O}$ (40 g, 0.15 mol), zinc granules [20 mesh] (18.3 g, 0.28 mol), mercury (13.55 g, 1 mL, 0.0676 mol) and 500 mL distilled water. Following flushing with N_2 for one hour, the mixture is warmed to about 80°C . to initiate the reaction. Following two hours of stirring under a constant N_2 flow, the mixture is cooled to room temperature and continuously stirred for another 48 hours by which time the reaction mixture turns into a deep blue solution. The solution is transferred into N_2 purged serum bottles and stored at 4°C . for future use.

[0197] The metal sulfide solution is prepared by adding about 950 mL Solution A into a 1 L fermenter and sparging with N_2 gas. An 85% solution of H_3PO_4 (1.5 mL, 30 mM) is added and the pH adjusted to 5.3 using concentrated aqueous NH_4OH . Solution B (10 mL) is added and the solution sparged with N_2 . Chromium (II) chloride is added until the oxidation-reduction potential (ORP) of the solution decreases to approximately -200 mV , wherein resazurin (1 mL of a 2 g/L solution) is added. Solution C (1/10; 10 ml FeCl_3 ; 5 ml CoCl_2 ; 5 ml NiCl_2 ; 1 ml H_3BO_3 ; 1 ml Na_2MoO_4 ; 1 ml MnCl_2 ; 1 ml Na_2WO_4 ; 1 ml ZnCl_2 ; 1 ml Na_2SeO_3 ; into 1 L media) is added, then sodium polysulfide (2 mL of a 3M solution) is added, and then the solution is sparged with N_2 gas.

[0198] Fermentation of a substrate comprising CO by *C. autoethanogenum* under batch conditions in the presence of polysulfide results in a substantially increased rate of accumulation and a final biomass accumulation of approximately 4 g/L over a 2-3 day period. For example, following a short lag phase of approximately 1 day, the biomass can increase from about 0.5 g/L up to at least 3.5 g/L over approximately 36 hours of fermentation. Furthermore, acetate is not produced during the growth phase in the presence of polysulfide (as is typically found in batch fermentations) and in certain circumstances some of the acetate is consumed, such that there is a net decrease in the amount of acetate in the fermenter. Culture biomass was harvested by centrifugation,

the cells washed once in media and cell pellets were either frozen at -80°C . or used immediately for fractionation of cellular components.

[0199] A semi-continuous fermentation approach can also be applied to maintain biomass productivity and reduce time associated with fermentation shut-down and start-up (i.e., turn-around time or lead time).

[0200] Harvesting of the accumulated bacterial biomass was performed at approximately 12-24 hour intervals, as the culture density approached (but before entering) stationary phase. Approximately half of the bioreactor volume was removed by transferring to a separate container via centrifugal pump. An equal volume of fresh or recycled media was then returned to the bioreactor such that the optical density of the reactor was approximately half of its initial value. The bioreactor fermentation was continued according to the above protocol so that multiple cycles of growth and biomass recovery was carried out during a single fermentation run.

[0201] b. *Clostridium ljungdahlii* Fermentation

[0202] Fermentation of *Clostridium ljungdahlii* is performed using similar methods to those described in, for example, U.S. Pat. Nos. 5,173,429 and 5,593,886. Briefly, batch fermentations are conducted using a biologically pure culture of *C. ljungdahlii*. Preparation of the medium ((1) 80.0 mL of a salt comprising KH_2PO_4 3.00 g/L, K_2HPO_4 3.00 g/L, $(\text{NH}_4)_2\text{SO}_4$ 6.00 g/L, NaCl 6.00 g/L, $\text{MgSO}_4 \cdot 2\text{H}_2\text{O}$ 1.25 g/L; (2) 1.0 g of yeast extract; (3) 1.0 g of trypticase; (4) 3.0 ml of PFN (Pfenning) trace metal solution comprising $\text{FeCl}_2 \cdot 4\text{H}_2\text{O}$ 1500 mg, $\text{ZnSO}_4 \cdot 7\text{H}_2\text{O}$ 100 mg, $\text{MnCl}_2 \cdot 4\text{H}_2\text{O}$ 30 mg, H_3BO_3 300 mg, $\text{CoCl}_2 \cdot 6\text{H}_2\text{O}$ 200 mg, $\text{CuCl}_2 \cdot \text{H}_2\text{O}$ 10 mg, $\text{NiCl}_2 \cdot 6\text{H}_2\text{O}$ 20 mg, $\text{NaMoO}_4 \cdot 2\text{H}_2\text{O}$ 30 mg, Na_2SeO_3 10 mg, and distilled water up to 1 L; (5) 10.0 ml of B vitamins comprising Pyridoxal HCl 10 mg, Riboflavin 50 mg, Thiamine HCl 50 mg, Nicotinic acid 50 mg, Ca-D-Pantothenate 50 mg, Lipoic acid 60 mg, p-aminobenzoic acid 50 mg, Folic acid 20 mg, Biotin 20 mg, cyanocobalamin 50 mg, and distilled water up to 1 L; (6) 0.5 g of cysteine HCl; (7) 0.06 g $\text{CaCl}_2 \cdot 2\text{H}_2\text{O}$; (8) 2.0 g NaHCO_3 ; (9) 1.0 mL resazurin (0.01%); and (10) 920.0 mL distilled water) is carried out anaerobically in an atmosphere of 80% nitrogen and 20% CO_2 . The pH of the medium is controlled during fermentation and maintained at 5.0 with HCl. If required, adjustments to the pH are made with sterile 10% NaOH or 1.0% acetic acid solution. The medium is transferred to 157.5 mL serum bottles and sealed with butyl rubber stoppers and aluminum seals. The bottles are then autoclaved at 121°C . for 20 minutes.

[0203] Approximately 48 hours before commencing the experiment, a seed culture is prepared from a stock culture of the *C. ljungdahlii* in a bottle similar to those as described above. The seed culture is grown in a shaker incubator at 37°C . and shaken at 100 rpm. Reducing solutions (2.0 ml Na_2S , 2.5% solution and 2.0 ml cysteine-HCl, 3.5% solution) are added to the culture, which is placed in the shaker incubator for approximately 15 minutes to allow for complete oxygen removal and temperature acclimation. Unlike the procedure used for isolating a biologically pure culture of the organism, addition of methane inhibitors is not required in batch fermentations.

[0204] Fermentation with *C. ljungdahlii* is performed in a New Brunswick Scientific Bioflow IIc 2.5-liter fermenter containing nutrient media at 37°C ., and a constant fluid level of 1.5 liters is maintained while the fluid is agitated at

variable rates of up to 1,000 revolutions per minute with gas introduced at a rate of approximately 500 cubic centimeters per minute. Optimal gas retention times are in the range of three minutes. The gas feed is varied with its uptake by the bacteria, which is in turn a function of the cell density.

[0205] Harvesting of the accumulated bacterial biomass was performed at approximately 12-24 hour intervals, as the culture density approached (but before entering) stationary phase. Approximately half of the bioreactor volume was removed by transferring to a separate container via centrifugal pump. An equal volume of fresh or recycled media was then returned to the bioreactor such that the optical density of the reactor was approximately half of its initial value. The bioreactor fermentation was continued according to the above protocol so that multiple cycles of growth and biomass recovery was carried out during a single fermentation run.

Example 2

Stable Carbon Isotope Distribution in Lipids from C_1 Metabolizing Microorganisms

[0206] Dry samples of *M. trichosporium* biomass and lipid fractions were analyzed for carbon and nitrogen content (% dry weight), and carbon (^{13}C) and nitrogen (^{15}N) stable isotope ratios via elemental analyzer/continuous flow isotope ratio mass spectrometry using a CHNOS Elemental Analyzer (vario ISOTOPE cube, Elementar, Hanau, Germany) coupled with an IsoPrime100 IRMS (Isoprime, Cheadle, UK). Samples of methanotrophic biomass cultured in fermenters or serum bottles were centrifuged, resuspended in deionized water and volumes corresponding to 0.2-2 mg carbon (about 0.5-5 mg dry cell weight) were transferred to 5x9 mm tin capsules (Costech Analytical Technologies, Inc., Valencia, Calif.) and dried at 80°C . for 24 hours. Similarly, previously extracted lipid fractions were suspended in chloroform and volumes containing 0.1-1.5 mg carbon were transferred to tin capsules and evaporated to dryness at 80°C . for 24 hours. Standards containing 0.1 mg carbon provided reliable $\delta^{13}\text{C}$ values.

[0207] The isotope ratio is expressed in “delta” notation (‰), wherein the isotopic composition of a material relative to that of a standard on a per million deviation basis is given by $\delta^{13}\text{C}$ (or $\delta^{15}\text{N}$) = $(R_{\text{Sample}}/R_{\text{Standard-1}}) \times 1,000$, wherein R is the molecular ratio of heavy to light isotope forms. The standard for carbon is the Vienna Pee Dee Belemnite (V-PDB) and for nitrogen is air. The NIST (National Institute of Standards and Technology) proposed SRM (Standard Reference Material) No. 1547, peach leaves, was used as a calibration standard. All isotope analyses were conducted at the Center for Stable Isotope Biogeochemistry at the University of California, Berkeley. Long-term external precision for C and N isotope analyses is 0.10‰ and 0.15‰, respectively.

[0208] *M. trichosporium* strain OB3b was grown on methane in three different fermentation batches, *M. capsulatus* Bath was grown on methane in two different fermentation batches, and *Methylomonas* sp. 16a was grown on methane in a single fermentation batch. The biomass from each of these cultures was analyzed for stable carbon isotope distribution ($\delta^{13}\text{C}$ values; see Table 3).

TABLE 3

Stable Carbon Isotope Distribution in Different Methanotrophs					
Methanotroph	Batch No.	EFT (h)†	OD ₆₀₀	DCW*	δ ¹³ C Cells
Mt OB3b	68A	48	1.80	1.00	-57.9
		64	1.97	1.10	-57.8
		71	2.10	1.17	-58.0
		88	3.10	1.73	-58.1
		97	4.30	2.40	-57.8
		113	6.00	3.35	-57.0
Mt OB3b	68B	127	8.40	4.69	-56.3
		32	2.90	1.62	-58.3
		41	4.60	2.57	-58.4
		47	5.89	3.29	-58.0
Mt OB3b	68C	56	7.90	4.41	-57.5
		72	5.32	2.97	-57.9
		79.5	5.90	3.29	-58.0
		88	5.60	3.12	-57.8
		94	5.62	3.14	-57.7
Mc Bath	62B	23	9.34	3.32	-60.8
		26	10.30	3.66	-60.1
		10	2.47	0.88	-59.9
		17.5	5.80	2.06	-61.0
		20	7.32	2.60	-61.1
Mc Bath	62A	23	9.34	3.32	-60.8
		26	10.30	3.66	-60.1
		10	2.95	1.05	-55.9
		13.5	3.59	1.27	-56.8
		17.5	5.40	1.92	-55.2
Mms 16a	66B	23	6.08	2.16	-57.2
		26	6.26	2.22	-57.6
		16	2.13	0.89	-65.5
		18	2.59	1.09	-65.1
		20.3	3.62	1.52	-65.5
27	5.50	2.31	-66.2		
40.5	9.80	4.12	-66.3		

*DCW, Dry Cell Weight is reported in g/L calculated from the measured optical densities (OD₆₀₀) using specific correlation factors relating OD of 1.0 to 0.358 g/L for Mt OB3b, OD of 1.0 to 0.355 g/L for Mc Bath, and OD of 1.0 to 0.42 g/L for Mms 16a. For Mt OB3b, the initial concentration of bicarbonate used per fermentation was 1.2 mM or 0.01% (Batch No. 68C) and 0.1% or 12 mM (Batch Nos. 68A and 68B).

†EFT = effective fermentation time in hours

[0209] In addition, stable carbon isotope analysis was performed for biomass and corresponding lipid fractions (see Table 4) from strains *Methylosinus trichosporium* OB3b (Mt OB3b), *Methylococcus capsulatus* Bath (Mc Bath), and *Methylomonas* sp. 16a (Mms 16a) grown on methane in bioreactors as described in Example 1.

TABLE 4

Stable Carbon Isotope Distribution in Cells and Lipids			
Batch No.	Strain	δ ¹³ C Cells	δ ¹³ C Lipids
68C	Mt OB3b	-57.7	-48.6
62A	Mc Bath	-57.6	-52.8
66A	Mms 16a	-64.4	-42.2

[0210] Biomass from strains Mt OB3b, Mc Bath and Mms 16a were harvested at 94 h (3.14 g DCW/L), 26 h (2.2 g DCW/L) and 39 h (1.14 g DCW/L), respectively. The δ¹³C values for lipids in Table 4 represent an average of duplicate determinations.

Example 3

Effect of Methane Source and Purity on Stable Carbon Isotope Distribution in Lipids

[0211] To examine methanotroph growth on methane containing natural gas components, a series of 0.5-liter serum bottles containing 100 mL defined media MMS1.0 were inoculated with *Methylosinus trichosporium* OB3b or *Meth-*

yllococcus capsulatus Bath from a serum bottle batch culture (5% v/v) grown in the same media supplied with a 1:1 (v/v) mixture of methane and air. The composition of medium MMS1.0 was as follows: 0.8 mM MgSO₄*7H₂O, 30 mM NaNO₃, 0.14 mM CaCl₂, 1.2 mM NaHCO₃, 2.35 mM KH₂PO₄, 3.4 mM K₂HPO₄, 20.7 μM Na₂MoO₄*2H₂O, 6 μM CuSO₄*5H₂O, 10 μM Fe^{III}-Na-EDTA, and 1 mL per liter of a trace metals solution (containing, per L: 500 mg FeSO₄*7H₂O, 400 mg ZnSO₄*7H₂O, 20 mg MnCl₂*7H₂O, 50 mg CoCl₂*6H₂O, 10 mg NiCl₂*6H₂O, 15 mg H₃BO₃, 250 mg EDTA). Phosphate, bicarbonate, and Fe^{III}-Na-EDTA were added after media was autoclaved and cooled. The final pH of the media was 7.0±0.1.

[0212] The inoculated bottles were sealed with rubber sleeve stoppers and injected with 60 mL methane gas added via syringe through sterile 0.45 μm filter and sterile 27G needles. Duplicate cultures were each injected with 60 mL volumes of (A) methane of 99% purity (grade 2.0, Praxair through Alliance Gas, San Carlos, Calif.), (B) methane of 70% purity representing a natural gas standard (Sigma-Aldrich; also containing 9% ethane, 6% propane, 3% methylpropane, 3% butane, and other minor hydrocarbon components), (C) methane of 85% purity delivered as a 1:1 mixture of methane sources A and B; and (D) >93% methane (grade 1.3, Specialty Chemical Products, South Houston, Tex.; in-house analysis showed composition >99% methane). The cultures were incubated at 30° C. (*M. trichosporium* strain OB3b) or 42° C. (*M. capsulatus* Bath) with rotary shaking at 250 rpm and growth was measured at approximately 12 hour intervals by withdrawing 1 mL samples to determine OD₆₀₀. At these times, the bottles were vented and headspace replaced with 60 mL of the respective methane source (A, B, C, or D) and 60 mL of concentrated oxygen (at least 85% purity). At about 24 hour intervals, 5 mL samples were removed, cells recovered by centrifugation (8,000 rpm, 10 minutes), and then stored at -80° C. before analysis.

[0213] Analysis of carbon and nitrogen content (% dry weight), and carbon (¹³C) and nitrogen (¹⁵N) stable isotope ratios, for methanotrophic biomass derived from *M. trichosporium* strain OB3b and *M. capsulatus* Bath were carried out. Table 5 shows the results of stable carbon isotope analysis for biomass samples from *M. capsulatus* Bath grown on methane having different levels of purity and in various batches of bottle cultures.

TABLE 5

Stable Carbon Isotope Distribution of <i>M. capsulatus</i> Bath Grown on Different Methane Sources having Different Purity					
Methane*	Batch No.	Time (h)†	OD ₆₀₀	DCW (g/L)	δ ¹³ C Cells
A	62C	22	1.02	0.36	-40.3
		56	2.01	0.71	-41.7
		73	2.31	0.82	-42.5
62D	62D	22	1.14	0.40	-39.3
		56	2.07	0.73	-41.6
		73	2.39	0.85	-42.0
B	62E	22	0.47	0.17	-44.7
		56	0.49	0.17	-45.4
		73	0.29	0.10	-45.4
62F	62F	22	0.62	0.22	-42.3
		56	0.63	0.22	-43.6
		73	0.30	0.11	-43.7

TABLE 5-continued

Stable Carbon Isotope Distribution of <i>M. capsulatus</i> Bath Grown on Different Methane Sources having Different Purity					
Methane*	Batch No.	Time (h)†	OD ₆₀₀	DCW (g/L)	δ ¹³ C Cells
C	62G	22	0.70	0.25	-40.7
		56	1.14	0.40	-44.8
		73	1.36	0.48	-45.8
	62H	22	0.62	0.22	-40.9
		56	1.03	0.37	-44.7
		73	1.23	0.44	-45.9

*Methane purity: A: 99% methane, grade 2.0 (min. 99%); B: 70% methane, natural gas standard (contains 9% ethane, 6% propane, 3% methylpropane, 3% butane); C: 85% methane (1:1 mix of A and B methane)

†Time = bottle culture time in hours

[0214] The average δ¹³C for *M. capsulatus* Bath grown on one source of methane (A, 99%) was -41.2±1.2, while the average δ¹³C for *M. capsulatus* Bath grown on a different source of methane (B, 70%) was -44.2±1.2. When methane sources A and B were mixed, an intermediate average δ¹³C of -43.8±2.4 was observed. These data show that the δ¹³C of cell material grown on methane sources A and B are significantly different from each other due to the differences in the δ¹³C of the input methane. But, cells grown on a mixture of the two gasses preferentially utilize ¹²C and, therefore, show a trend to more negative δ¹³C values.

[0215] A similar experiment was performed to examine whether two different methanotrophs, *Methylococcus capsulatus* Bath and *Methylosinus trichosporium* OB3b, grown on different methane sources and in various batches of bottle cultures showed a difference in δ¹³C distribution (see Table 6).

TABLE 6

Stable Carbon Isotope Distribution of Different Methanotrophs Grown on Different Methane Sources of Different Purity						
Strain	Methane*	Batch No.	Time (h)†	OD ₆₀₀	DCW (g/L)	δ ¹³ C Cells
Mc Bath	A	62I	18	0.494	0.18	-54.3
			40	2.33	0.83	-42.1
			48	3.08	1.09	-37.1
Mc Bath	D	62J	18	0.592	0.21	-38.3
			40	1.93	0.69	-37.8
			48	2.5	0.89	-37.8
Mc Bath	D	62K	18	0.564	0.20	-38.6
			40	1.53	0.54	-37.5
			48	2.19	0.78	-37.6
Mt OB3b	A	68D	118	0.422	0.24	-50.2
			137	0.99	0.55	-47.7
			162	1.43	0.80	-45.9
Mt OB3b	A	68E	118	0.474	0.26	-49.9
			137	1.065	0.59	-47.6
			162	1.51	0.84	-45.2
Mt OB3b	D	68F	118	0.534	0.30	-45.6
			137	1.119	0.62	-38.7
			162	1.63	0.91	-36.4
Mt OB3b	D	68G	118	0.544	0.30	-44.8
			137	1.131	0.63	-39.1
			162	1.6	0.89	-34.2

*Methane sources and purity:

A: 99% methane (grade 2.0);

D: >93% methane (grade 1.3)

†Time = bottle culture time in hours

[0216] The average δ¹³C for *M. capsulatus* grown on a first methane source (A) was -44.5±8.8, while the average δ¹³C for *M. trichosporium* was -47.8±2.0 grown on the same methane source. The average δ¹³C for *M. capsulatus* grown on the second methane source (B) was -37.9±0.4, while the average δ¹³C for *M. trichosporium* was -39.8±4.5. These data show that the δ¹³C of cell material grown on a methane source is highly similar to the δ¹³C of cell material from a different strain grown on the same source of methane. Thus, the observed δ¹³C of cell material appears to be primarily dependent on the composition of the input gas rather than a property of a particular bacterial strain being studied.

[0217] The various embodiments described above can be combined to provide further embodiments. All of the patent and non-patent publications referred to in this specification or listed in the Application Data Sheet, including the disclosure of U.S. provisional application No. 61/928,366, filed Jan. 16, 2014, are incorporated herein by reference in their entirety. Aspects of the embodiments can be modified, if necessary to employ concepts of the various patents, applications and publications to provide further embodiments.

[0218] These and other changes can be made to the embodiments in light of the above-detailed description. In general, in the following embodiments, the terms used should not be construed to limit the embodiments to the specific embodiments disclosed in the specification and the embodiments, but should be construed to include all possible embodiments along with the full scope of equivalents to which such embodiments are entitled. Accordingly, the embodiments are not limited by the disclosure.

SEQUENCE LISTING

<160> NUMBER OF SEQ ID NOS: 38

<210> SEQ ID NO 1

<211> LENGTH: 852

<212> TYPE: DNA

<213> ORGANISM: Artificial sequence

<220> FEATURE:

<223> OTHER INFORMATION: Synthetic construct

<220> FEATURE:

<221> NAME/KEY: CDS

<222> LOCATION: (1)..(852)

<223> OTHER INFORMATION: *Saccharomyces cerevisiae* mature KRE1 protein
(codon optimized)

<400> SEQUENCE: 1

```

gtc atg gcc gct gtt acc acc caa gtc acc gtc gtc acc aat gtt gct      48
Val Met Ala Ala Val Thr Thr Gln Val Thr Val Val Thr Asn Val Ala
1          5          10          15

gga gcc ctt gtc acc gaa acc acc atc tgg gac ccc gcg acc gcc gca      96
Gly Ala Leu Val Thr Glu Thr Thr Ile Trp Asp Pro Ala Thr Ala Ala
20         25         30

gcc gca gcg acg acc acc gcc cag acg ggc ttc ttc acc acc gtg ttc      144
Ala Ala Ala Thr Thr Thr Ala Gln Thr Gly Phe Phe Thr Thr Val Phe
35         40         45

acg acc acc aac gac gtg ggt acc act gtc acg ctg acg cag acc gtg      192
Thr Thr Thr Asn Asp Val Gly Thr Thr Val Thr Leu Thr Gln Thr Val
50         55         60

aat cgc gcc acg atg ctg ccc act acc acg acg agc acc agc agc acc      240
Asn Arg Ala Thr Met Leu Pro Thr Thr Thr Thr Ser Thr Ser Thr Thr
65         70         75         80

ggc aag acc acc acg acg gtg ccg act gcg acg tcc tcg ctc agt tcg      288
Gly Lys Thr Thr Thr Val Pro Thr Ala Thr Ser Ser Leu Ser Ser
85         90         95

ggc ctg tcg acc gtc acc acc acg aac gac ctg ggc acg acc gtg acg      336
Gly Leu Ser Thr Val Thr Thr Thr Asn Asp Leu Gly Thr Thr Val Thr
100        105        110

ctc acc cag acc ttc acg cac tcc tcc acg tcc gcg acc tcc tcg gcc      384
Leu Thr Gln Thr Phe Thr His Ser Ser Thr Ser Ala Thr Ser Ser Ala
115        120        125

tcc tcg tcg gtg agc tcg tcc gtc agt agc agt ggc tcc tcc agc agc      432
Ser Ser Ser Val Ser Ser Ser Val Ser Ser Ser Gly Ser Ser Ser Ser
130        135        140

gtc aag acc acg acc tcg acc ggc tcc gcg gtg gcg gaa acc ggg tgg      480
Val Lys Thr Thr Thr Ser Thr Gly Ser Ala Val Ala Glu Thr Gly Trp
145        150        155        160

gac ccg agc acg gat ttt acc gag ccg cca gtg agc gcg gtc acc agc      528
Asp Pro Ser Thr Asp Phe Thr Glu Pro Pro Val Ser Ala Val Thr Ser
165        170        175

ctc tcg atc gac tcg tat ata acg atc acc gag ggc acc acc tcg acc      576
Leu Ser Ile Asp Ser Tyr Ile Thr Ile Thr Glu Gly Thr Thr Ser Thr
180        185        190

tac act acc acc cgg gcg ccg acc tcg atg tgg gtg acc gtc gtc cgc      624
Tyr Thr Thr Thr Arg Ala Pro Thr Ser Met Trp Val Thr Val Val Arg
195        200        205

cag ggg aac acg atc acc gtg caa acc acc ttc gtc cag cgc ttc agc      672
Gln Gly Asn Thr Ile Thr Val Gln Thr Thr Phe Val Gln Arg Phe Ser
210        215        220

tcc caa tac gtg acc gtg gat tcc gtc ggc agc atc ggc atg ggt acg      720
Ser Gln Tyr Val Thr Val Asp Ser Val Gly Ser Ile Gly Met Gly Thr
225        230        235        240

```

-continued

```

ctg acc ggt acc gtc ggc gtg atc aag tcc gcc atc aag aaa acc gtg      768
Leu Thr Gly Thr Val Gly Val Ile Lys Ser Ala Ile Lys Lys Thr Val
                245                      250                      255

tcc cat aac gag gcc cag cat ctc ggc atg tcg tcg ttc acg tcg att      816
Ser His Asn Glu Ala Gln His Leu Gly Met Ser Ser Phe Thr Ser Ile
                260                      265                      270

ctg ggt ggc ctc ctc acg gtc ttg atc tgg ttc ctg      852
Leu Gly Gly Leu Leu Thr Val Leu Ile Trp Phe Leu
                275                      280

<210> SEQ ID NO 2
<211> LENGTH: 284
<212> TYPE: PRT
<213> ORGANISM: Artificial sequence
<220> FEATURE:
<223> OTHER INFORMATION: Synthetic Construct

<400> SEQUENCE: 2
Val Met Ala Ala Val Thr Thr Gln Val Thr Val Val Thr Asn Val Ala
1          5          10          15
Gly Ala Leu Val Thr Glu Thr Thr Ile Trp Asp Pro Ala Thr Ala Ala
20         25         30
Ala Ala Ala Thr Thr Thr Ala Gln Thr Gly Phe Phe Thr Thr Val Phe
35         40         45
Thr Thr Thr Asn Asp Val Gly Thr Thr Val Thr Leu Thr Gln Thr Val
50         55         60
Asn Arg Ala Thr Met Leu Pro Thr Thr Thr Thr Ser Thr Ser Ser Thr
65         70         75         80
Gly Lys Thr Thr Thr Thr Val Pro Thr Ala Thr Ser Ser Leu Ser Ser
85         90         95
Gly Leu Ser Thr Val Thr Thr Thr Asn Asp Leu Gly Thr Thr Val Thr
100        105        110
Leu Thr Gln Thr Phe Thr His Ser Ser Thr Ser Ala Thr Ser Ser Ala
115        120        125
Ser Ser Ser Val Ser Ser Ser Val Ser Ser Ser Gly Ser Ser Ser Ser
130        135        140
Val Lys Thr Thr Thr Ser Thr Gly Ser Ala Val Ala Glu Thr Gly Trp
145        150        155        160
Asp Pro Ser Thr Asp Phe Thr Glu Pro Pro Val Ser Ala Val Thr Ser
165        170        175
Leu Ser Ile Asp Ser Tyr Ile Thr Ile Thr Glu Gly Thr Thr Ser Thr
180        185        190
Tyr Thr Thr Thr Arg Ala Pro Thr Ser Met Trp Val Thr Val Val Arg
195        200        205
Gln Gly Asn Thr Ile Thr Val Gln Thr Thr Phe Val Gln Arg Phe Ser
210        215        220
Ser Gln Tyr Val Thr Val Asp Ser Val Gly Ser Ile Gly Met Gly Thr
225        230        235        240
Leu Thr Gly Thr Val Gly Val Ile Lys Ser Ala Ile Lys Lys Thr Val
245        250        255
Ser His Asn Glu Ala Gln His Leu Gly Met Ser Ser Phe Thr Ser Ile
260        265        270
Leu Gly Gly Leu Leu Thr Val Leu Ile Trp Phe Leu
275        280

```

-continued

```

<210> SEQ ID NO 3
<211> LENGTH: 4065
<212> TYPE: DNA
<213> ORGANISM: Artificial sequence
<220> FEATURE:
<223> OTHER INFORMATION: Synthetic construct
<220> FEATURE:
<221> NAME/KEY: CDS
<222> LOCATION: (1)..(4065)
<223> OTHER INFORMATION: Saccharomyces cerevisiae mature KRE2 protein
(codon optimized)

<400> SEQUENCE: 3

atg cgc ctt ctt gcc ctt gtc ttg ttg ctt ctc tgt gca ccg ctt cgc      48
Met Arg Leu Leu Ala Leu Val Leu Leu Leu Leu Cys Ala Pro Leu Arg
1          5          10          15

gcc tgg acc tat tcg ctc cgc tat ggt atc ccc gag tcc gcc cag gtg      96
Ala Trp Thr Tyr Ser Leu Arg Tyr Gly Ile Pro Glu Ser Ala Gln Val
          20          25          30

tgg tcg atc ctc gtt cat ctg ctc ggc gac gtg gac aac caa ctc ctt     144
Trp Ser Ile Leu Val His Leu Leu Gly Asp Val Asp Asn Gln Leu Leu
          35          40          45

act aac ctg tat ccc ctg gtg acc ggg ctc gac gat gag atc gac atc     192
Thr Asn Leu Tyr Pro Leu Val Thr Gly Leu Asp Asp Glu Ile Asp Ile
          50          55          60

cag gag aac ctc gtt acg tcc aac gtg ctg cgc gag cgc tac gat aaa     240
Gln Glu Asn Leu Val Thr Ser Asn Val Leu Arg Glu Arg Tyr Asp Lys
          65          70          75          80

gag gac gtc gcg gat ctg ctg gaa ctc tac gca tcg ctc tac ccc atg     288
Glu Asp Val Ala Asp Leu Leu Glu Leu Tyr Ala Ser Leu Tyr Pro Met
          85          90          95

ggg atg atc caa cac gac atc tcg tcg aat gcc gag caa gac gac gcg     336
Gly Met Ile Gln His Asp Ile Ser Ser Asn Ala Glu Gln Asp Asp Ala
          100          105          110

aat tcc tcc tat ttc gtc ctg aac ggc aat cgg tat gag aaa ccc gac     384
Asn Ser Ser Tyr Phe Val Leu Asn Gly Asn Arg Tyr Glu Lys Pro Asp
          115          120          125

gac gtc ttt tac ctg aag tcg aag gac ctg acc atc cag cag aaa gtg     432
Asp Val Phe Tyr Leu Lys Ser Lys Asp Leu Thr Ile Gln Gln Lys Val
          130          135          140

ccc gat gtc gac gtc atc caa ccg tac gac gtc gtg att gga acc aac     480
Pro Asp Val Asp Val Ile Gln Pro Tyr Asp Val Val Ile Gly Thr Asn
          145          150          155          160

tcc gag gcg ccc ata ctg atc ctc tac ggc tgc ccg acc gtc atc gac     528
Ser Glu Ala Pro Ile Leu Ile Leu Tyr Gly Cys Pro Thr Val Ile Asp
          165          170          175

agc gac ttc gag gag ttc aat cgg aat ctc ttc atg gag gct atg aac     576
Ser Asp Phe Glu Glu Phe Asn Arg Asn Leu Phe Met Glu Ala Met Asn
          180          185          190

ggc gag gcc aag ttc cgc ttc att tgg cgg agc acg tgt agc ctg gac     624
Gly Glu Gly Lys Phe Arg Phe Ile Trp Arg Ser Thr Cys Ser Leu Asp
          195          200          205

ggc aag tcc gtg gag tac ccg ctg acc cac ccc ctg gag atc acc ctg     672
Gly Lys Ser Val Glu Tyr Pro Leu Thr His Pro Leu Glu Ile Thr Leu
          210          215          220

cag aac gcc agc cgc atg tcc tcc atc cct cag ctc aag aag atc ctg     720
Gln Asn Gly Ser Arg Met Ser Ser Ile Pro Gln Leu Lys Lys Ile Leu
          225          230          235          240

```

-continued

tac acc gtt ccg aaa gaa atc ctc gtg ggc gca gat aac gac gac cag	768
Tyr Thr Val Pro Lys Glu Ile Leu Val Gly Ala Asp Asn Asp Asp Gln	
245 250 255	
ctg cac gac ctg gag ccc gag gag ctg cgc gag ctg gac ctc cgc gtg	816
Leu His Asp Leu Glu Pro Glu Glu Leu Arg Glu Leu Asp Leu Arg Val	
260 265 270	
acc tcg ctc att tcc gag ttc tat cag tat aag aaa gac att acg gcc	864
Thr Ser Leu Ile Ser Glu Phe Tyr Gln Tyr Lys Lys Asp Ile Thr Ala	
275 280 285	
acc ctg aat ttc acc aaa agt atc gtc aac aat ttc ccg ctg att tcg	912
Thr Leu Asn Phe Thr Lys Ser Ile Val Asn Asn Phe Pro Leu Ile Ser	
290 295 300	
aag cag ctg atc aag gtt tcg tcg gtc aat aaa gac atc atc acc tcc	960
Lys Gln Leu Ile Lys Val Ser Ser Val Asn Lys Asp Ile Ile Thr Ser	
305 310 315 320	
aac gag gag ttg aat tcc aag ggc ttc gac tac aac atg ctg ggc atc	1008
Asn Glu Glu Leu Asn Ser Lys Gly Phe Asp Tyr Asn Met Leu Gly Ile	
325 330 335	
aac ggc cag aac tgg aag atc acc tcc ctg acg ccc tac aat ctt ctc	1056
Asn Gly Gln Asn Trp Lys Ile Thr Ser Leu Thr Pro Tyr Asn Leu Leu	
340 345 350	
acg gcc ctg aaa acg gag tac cag agt ctg ctg aag atc acc aac ctc	1104
Thr Ala Leu Lys Thr Glu Tyr Gln Ser Leu Leu Lys Ile Thr Asn Leu	
355 360 365	
ctc cag gag ctg gag ccc tcc aag tgc atc ctc gac tcc aag ttc ctg	1152
Leu Gln Glu Leu Glu Pro Ser Lys Cys Ile Leu Asp Ser Lys Phe Leu	
370 375 380	
ctc aat aag ttc tcg cag ttc agc ctg ggt aaa ctg cag aat ctg caa	1200
Leu Asn Lys Phe Ser Gln Phe Ser Leu Gly Lys Leu Gln Asn Leu Gln	
385 390 395 400	
ccg atc aaa atg gac ctc cat acc atc ccg ggt ttt agc gag tcc gtc	1248
Pro Ile Lys Met Asp Leu His Thr Ile Pro Gly Phe Ser Glu Ser Val	
405 410 415	
atc tac ttc aat gat att gag agt gac ccg cag tac gac gag ctc gtc	1296
Ile Tyr Phe Asn Asp Ile Glu Ser Asp Pro Gln Tyr Asp Glu Leu Val	
420 425 430	
aac tcg gtg caa gca ttc ttc gac aag tcg aag ttc ggc gag ctg ccc	1344
Asn Ser Val Gln Ala Phe Phe Asp Lys Ser Lys Phe Gly Glu Leu Pro	
435 440 445	
gag atc aag cag aac tgg tcc gag att atc ttc gtc ata gat ttt gcc	1392
Glu Ile Lys Gln Asn Trp Ser Glu Ile Ile Phe Val Ile Asp Phe Ala	
450 455 460	
cgg ctg gag gac tcg gaa gtc aaa gag gcc ctc ggc gga ctg gtg agg	1440
Arg Leu Glu Asp Ser Glu Val Lys Glu Ala Leu Gly Gly Leu Val Arg	
465 470 475 480	
gct gtg aac gtc gtg agc cag ggg tac ccc cag cgc gtg gga ctc ctc	1488
Ala Val Asn Val Val Ser Gln Gly Tyr Pro Gln Arg Val Gly Leu Leu	
485 490 495	
ccg ttc agc agt gat agc gac aag agc gtc gtc aat aag atc tac gag	1536
Pro Phe Ser Ser Asp Ser Asp Lys Ser Val Val Asn Lys Ile Tyr Glu	
500 505 510	
ctg aag aac tcg acc gac aat ctc acc gag ctg aag tcg ttc ctg gaa	1584
Leu Lys Asn Ser Thr Asp Asn Leu Thr Glu Leu Lys Ser Phe Leu Glu	
515 520 525	
acc atg ttg ctg gcc gac ggc ctg tcc gcc aac gcg aag cat agt aag	1632
Thr Met Leu Leu Ala Asp Gly Leu Ser Ala Asn Ala Lys His Ser Lys	
530 535 540	

-continued

cat atc ccc gtg ccg gac gtg ttc cac ctc ctc gac gag ctg cag atc	1680
His Ile Pro Val Pro Asp Val Phe His Leu Leu Asp Glu Leu Gln Ile	
545 550 555 560	
gac gaa acg tcc atc atc atc aac ggc gag ata tac ccg ttc cgc aag	1728
Asp Glu Thr Ser Ile Ile Ile Asn Gly Glu Ile Tyr Pro Phe Arg Lys	
565 570 575	
aat tgg aac tac ctc atc gcc aag gtc atc aag aaa gac acc gaa ttc	1776
Asn Trp Asn Tyr Leu Ile Ala Lys Val Ile Lys Lys Asp Thr Glu Phe	
580 585 590	
atc cgc aag gag ctg tcg aac tcg tcg ccg aag aac aag cag att agt	1824
Ile Arg Lys Glu Leu Ser Asn Ser Ser Pro Lys Asn Lys Gln Ile Ser	
595 600 605	
gtg cgc gac ctg ttg cac tat aag agc gcg aac ctc cgc cat aac aag	1872
Val Arg Asp Leu Leu His Tyr Lys Ser Ala Asn Leu Arg His Asn Lys	
610 615 620	
tat acg ccg aac tat ttc gcg gat agt gtg tat tcc tcg gtc aac aat	1920
Tyr Thr Pro Asn Tyr Phe Ala Asp Ser Val Tyr Ser Ser Val Asn Asn	
625 630 635 640	
acc gct ctg gaa agc gtc tgc tcg atc ggt tac tac acc aaa aac gag	1968
Thr Ala Leu Glu Ser Val Cys Ser Ile Gly Tyr Tyr Thr Lys Asn Glu	
645 650 655	
gaa tat aac ctc ctg cat acc att acg ctc gtg gat gac ttc ggc tcg	2016
Glu Tyr Asn Leu Leu His Thr Ile Thr Leu Val Asp Asp Phe Gly Ser	
660 665 670	
atc cat gcg ctg aag cgg ctg cgg aac ctg ttg cat acg tcc ttc gtg	2064
Ile His Ala Leu Lys Arg Leu Arg Asn Leu Leu His Thr Ser Phe Val	
675 680 685	
ggc gtg cgg atc cgc att atc cat gtc ggc gat atc agc gac atc tgg	2112
Gly Val Arg Ile Arg Ile Ile His Val Gly Asp Ile Ser Asp Ile Trp	
690 695 700	
tat cag ctc cgc gga tcc ctg agt cag aaa gac ccg atc ggc agc atc	2160
Tyr Gln Leu Arg Gly Ser Leu Ser Gln Lys Asp Pro Ile Gly Ser Ile	
705 710 715 720	
aac acc ttc atc gac gcc ctg aaa ctc aaa aag gtc aag tcc cat acg	2208
Asn Thr Phe Ile Asp Ala Leu Lys Leu Lys Lys Val Lys Ser His Thr	
725 730 735	
tat aag aag tcg cag cag ctc ggc ttg cat aag tgg ctc ccc gac atc	2256
Tyr Lys Lys Ser Gln Gln Leu Gly Leu His Lys Trp Leu Pro Asp Ile	
740 745 750	
ccg ctg ttc gag ctc caa aag ggt tcg ttc atc gcg ctc aac ggc cgg	2304
Pro Leu Phe Glu Leu Gln Lys Gly Ser Phe Ile Ala Leu Asn Gly Arg	
755 760 765	
ttc atc atc ctg atc aag atg aaa tgc cag aag caa aac atc tcc aaa	2352
Phe Ile Ile Leu Ile Lys Met Lys Cys Gln Lys Gln Asn Ile Ser Lys	
770 775 780	
gcc aag atc atc aag cgc gag gcc ctt cgg acc ata gat tcg gtg ttc	2400
Ala Lys Ile Ile Lys Arg Glu Ala Leu Arg Thr Ile Asp Ser Val Phe	
785 790 795 800	
gcg ctg gac ctc ctc ttt cct ggc ttc agc caa gag atc ata aat ccc	2448
Ala Leu Asp Leu Leu Phe Pro Gly Phe Ser Gln Glu Ile Ile Asn Pro	
805 810 815	
gat ctc atc gag atg atc tcc tcg atc ctt acc cgc ctc ttc tat cag	2496
Asp Leu Ile Glu Met Ile Ser Ser Ile Leu Thr Arg Leu Phe Tyr Gln	
820 825 830	
ggg acc cac ata tac aac aac ggc att gac tat act acc gag tcg tcg	2544
Gly Thr His Ile Tyr Asn Asn Gly Ile Asp Tyr Thr Thr Glu Ser Ser	
835 840 845	

-continued

ctg ccg cgc atg gac ttg tcc gag ttc ttc cgc ccg aat aac ctg acc	2592
Leu Pro Arg Met Asp Leu Ser Glu Phe Phe Arg Pro Asn Asn Leu Thr	
850 855 860	
atg ttc gag gat ggc aaa tcg gcg tcc atc gat ctc ctc ctc atc ctt	2640
Met Phe Glu Asp Gly Lys Ser Ala Ser Ile Asp Leu Leu Leu Ile Leu	
865 870 875 880	
gac ccg ctg gaa gaa cgg act cag atg att ctt tcc ctc gtg gag caa	2688
Asp Pro Leu Glu Glu Arg Thr Gln Met Ile Leu Ser Leu Val Glu Gln	
885 890 895	
ttc cgg cca ctg aag ttc gtg aat atc cag gtc atc ctg atg ccg acc	2736
Phe Arg Pro Leu Lys Phe Val Asn Ile Gln Val Ile Leu Met Pro Thr	
900 905 910	
ctg gag ctg aat att gtc ccg atc cgg cgc atc tac gtg gac gac gcg	2784
Leu Glu Leu Asn Ile Val Pro Ile Arg Arg Ile Tyr Val Asp Asp Ala	
915 920 925	
gat atc gtc aag tcc atc acg tcc gag gac tcc cgg tcg gac cct gag	2832
Asp Ile Val Lys Ser Ile Thr Ser Glu Asp Ser Arg Ser Asp Pro Glu	
930 935 940	
gtt gac atc gag atg gat gtg ccg aac tcg ttc atc gtc gac aat aac	2880
Val Asp Ile Glu Met Asp Val Pro Asn Ser Phe Ile Val Asp Asn Asn	
945 950 955 960	
tac agg att aag aaa ctg ttg att gag ctg cat tcg ttc tcc agt aaa	2928
Tyr Arg Ile Lys Lys Leu Leu Ile Glu Leu His Ser Phe Ser Ser Lys	
965 970 975	
acc gtg ctg tcc acg ggc aat atc gac ggc atg ggt ggc gtg tgc ctt	2976
Thr Val Leu Ser Thr Gly Asn Ile Asp Gly Met Gly Gly Val Cys Leu	
980 985 990	
gcg ctc gtc gat tcg gct ggc aac att atc gac aaa acc acg acc atg	3024
Ala Leu Val Asp Ser Ala Gly Asn Ile Ile Asp Lys Thr Thr Thr Met	
995 1000 1005	
aaa acg ttc ggg tac ggc cag ttc cac acc gac aag ttc ttg aag	3069
Lys Thr Phe Gly Tyr Gly Gln Phe His Thr Asp Lys Phe Leu Lys	
1010 1015 1020	
ggt tgc tac atc aaa agc tgc gac agc cgc tat acc gtc cag tcc	3114
Gly Cys Tyr Ile Lys Ser Cys Asp Ser Arg Tyr Thr Val Gln Ser	
1025 1030 1035	
ttc agc act gat ggc cac ccc gat ttc atc ccg tcc gac tcc ctc	3159
Phe Ser Thr Asp Gly His Pro Asp Phe Ile Pro Ser Asp Ser Leu	
1040 1045 1050	
gac atc ctg agc tat aac ccg cag aag att gcg gtt aag atc tcc	3204
Asp Ile Leu Ser Tyr Asn Pro Gln Lys Ile Ala Val Lys Ile Ser	
1055 1060 1065	
gag gag ccg acg cac gaa gaa gag tat gaa gag ggt cgc aat aac	3249
Glu Glu Pro Thr His Glu Glu Glu Tyr Glu Glu Gly Arg Asn Asn	
1070 1075 1080	
gac acg atc atc aat atc ttt acc att tcg ggt ccc gat gaa gaa	3294
Asp Thr Ile Ile Asn Ile Phe Thr Ile Ser Gly Pro Asp Glu Glu	
1085 1090 1095	
gag agg tac atg caa atg atc ctg tcg atc ctc agc aaa tgc cca	3339
Glu Arg Tyr Met Gln Met Ile Leu Ser Ile Leu Ser Lys Cys Pro	
1100 1105 1110	
gaa acc caa aaa gtg aat ttc ttt ata ctg gac cag ccg ttc att	3384
Glu Thr Gln Lys Val Asn Phe Phe Ile Leu Asp Gln Pro Phe Ile	
1115 1120 1125	
tcg gac acc ctg cgc aag tcc tgc gaa tac atc aat tcc agt gat	3429
Ser Asp Thr Leu Arg Lys Ser Cys Glu Tyr Ile Asn Ser Ser Asp	
1130 1135 1140	

-continued

gag atg cgc ggc aat gtc ata ttc ctc aac tat gag tgg ccc cag	3474
Glu Met Arg Gly Asn Val Ile Phe Leu Asn Tyr Glu Trp Pro Gln	
1145 1150 1155	
tgg ctg cgc cct cag cgg ttc agc agc cgc agg cgc gac gtc agc	3519
Trp Leu Arg Pro Gln Arg Phe Ser Ser Arg Arg Arg Asp Val Ser	
1160 1165 1170	
cgg ttc ctc ttc ctg gac gtg ctc ctc ccg cag aac atc agc aag	3564
Arg Phe Leu Phe Leu Asp Val Leu Leu Pro Gln Asn Ile Ser Lys	
1175 1180 1185	
gtc ctc tat atg tcg ccg acc gag gtc ccg ctt gat ccg ttc gac	3609
Val Leu Tyr Met Ser Pro Thr Glu Val Pro Leu Asp Pro Phe Asp	
1190 1195 1200	
atc ttc caa ttc cag ggc ctc aag cgc gca ccg ctg ggc ctg ttt	3654
Ile Phe Gln Phe Gln Gly Leu Lys Arg Ala Pro Leu Gly Leu Phe	
1205 1210 1215	
cgc atg agc ggt gac ggc tat tgg aaa gag ggc tac tgg gag aag	3699
Arg Met Ser Gly Asp Gly Tyr Trp Lys Glu Gly Tyr Trp Glu Lys	
1220 1225 1230	
atg ctg cgc gag aac aac ttg gaa ttc tat agc acc gag ccg gcg	3744
Met Leu Arg Glu Asn Asn Leu Glu Phe Tyr Ser Thr Glu Pro Ala	
1235 1240 1245	
ttc ttg gtc aat ctg gaa cgc ttc cgc gag ctg gac gcc ggc gac	3789
Phe Leu Val Asn Leu Glu Arg Phe Arg Glu Leu Asp Ala Gly Asp	
1250 1255 1260	
aag tac agg atc cat tac caa cgc att agc acc gac gcg atg agc	3834
Lys Tyr Arg Ile His Tyr Gln Arg Ile Ser Thr Asp Ala Met Ser	
1265 1270 1275	
ctg gtg aac atc ggg caa gac ctc gtc aat aat ctg caa ctt gag	3879
Leu Val Asn Ile Gly Gln Asp Leu Val Asn Asn Leu Gln Leu Glu	
1280 1285 1290	
gtc ccg atc cgg ttc ctg aag ggt agt tat aag aaa aag ctc gtg	3924
Val Pro Ile Arg Phe Leu Lys Gly Ser Tyr Lys Lys Lys Leu Val	
1295 1300 1305	
atc aat gat gag tgc gtc agc gag tgg aag aaa aag atc aac aag	3969
Ile Asn Asp Glu Cys Val Ser Glu Trp Lys Lys Lys Ile Asn Lys	
1310 1315 1320	
ttt gcc tcg tcc cca ggg gac gag gac gtt ccc gcc gag agt gtg	4014
Phe Ala Ser Ser Pro Gly Asp Glu Asp Val Pro Gly Glu Ser Val	
1325 1330 1335	
agc tcg aag tat cag gat tcg gat aac gcc gcg cca ctc cat gac	4059
Ser Ser Lys Tyr Gln Asp Ser Asp Asn Ala Ala Pro Leu His Asp	
1340 1345 1350	
gaa ctc	4065
Glu Leu	
1355	

<210> SEQ ID NO 4
 <211> LENGTH: 1355
 <212> TYPE: PRT
 <213> ORGANISM: Artificial sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Synthetic Construct

<400> SEQUENCE: 4

Met Arg Leu Leu Ala Leu Val Leu Leu Leu Leu Cys Ala Pro Leu Arg
 1 5 10 15
 Ala Trp Thr Tyr Ser Leu Arg Tyr Gly Ile Pro Glu Ser Ala Gln Val
 20 25 30

-continued

Trp Ser Ile Leu Val His Leu Leu Gly Asp Val Asp Asn Gln Leu Leu
 35 40 45
 Thr Asn Leu Tyr Pro Leu Val Thr Gly Leu Asp Asp Glu Ile Asp Ile
 50 55 60
 Gln Glu Asn Leu Val Thr Ser Asn Val Leu Arg Glu Arg Tyr Asp Lys
 65 70 75 80
 Glu Asp Val Ala Asp Leu Leu Glu Leu Tyr Ala Ser Leu Tyr Pro Met
 85 90 95
 Gly Met Ile Gln His Asp Ile Ser Ser Asn Ala Glu Gln Asp Asp Ala
 100 105 110
 Asn Ser Ser Tyr Phe Val Leu Asn Gly Asn Arg Tyr Glu Lys Pro Asp
 115 120 125
 Asp Val Phe Tyr Leu Lys Ser Lys Asp Leu Thr Ile Gln Gln Lys Val
 130 135 140
 Pro Asp Val Asp Val Ile Gln Pro Tyr Asp Val Val Ile Gly Thr Asn
 145 150 155 160
 Ser Glu Ala Pro Ile Leu Ile Leu Tyr Gly Cys Pro Thr Val Ile Asp
 165 170 175
 Ser Asp Phe Glu Glu Phe Asn Arg Asn Leu Phe Met Glu Ala Met Asn
 180 185 190
 Gly Glu Gly Lys Phe Arg Phe Ile Trp Arg Ser Thr Cys Ser Leu Asp
 195 200 205
 Gly Lys Ser Val Glu Tyr Pro Leu Thr His Pro Leu Glu Ile Thr Leu
 210 215 220
 Gln Asn Gly Ser Arg Met Ser Ser Ile Pro Gln Leu Lys Lys Ile Leu
 225 230 235 240
 Tyr Thr Val Pro Lys Glu Ile Leu Val Gly Ala Asp Asn Asp Asp Gln
 245 250 255
 Leu His Asp Leu Glu Pro Glu Glu Leu Arg Glu Leu Asp Leu Arg Val
 260 265 270
 Thr Ser Leu Ile Ser Glu Phe Tyr Gln Tyr Lys Lys Asp Ile Thr Ala
 275 280 285
 Thr Leu Asn Phe Thr Lys Ser Ile Val Asn Asn Phe Pro Leu Ile Ser
 290 295 300
 Lys Gln Leu Ile Lys Val Ser Ser Val Asn Lys Asp Ile Ile Thr Ser
 305 310 315 320
 Asn Glu Glu Leu Asn Ser Lys Gly Phe Asp Tyr Asn Met Leu Gly Ile
 325 330 335
 Asn Gly Gln Asn Trp Lys Ile Thr Ser Leu Thr Pro Tyr Asn Leu Leu
 340 345 350
 Thr Ala Leu Lys Thr Glu Tyr Gln Ser Leu Leu Lys Ile Thr Asn Leu
 355 360 365
 Leu Gln Glu Leu Glu Pro Ser Lys Cys Ile Leu Asp Ser Lys Phe Leu
 370 375 380
 Leu Asn Lys Phe Ser Gln Phe Ser Leu Gly Lys Leu Gln Asn Leu Gln
 385 390 395 400
 Pro Ile Lys Met Asp Leu His Thr Ile Pro Gly Phe Ser Glu Ser Val
 405 410 415
 Ile Tyr Phe Asn Asp Ile Glu Ser Asp Pro Gln Tyr Asp Glu Leu Val
 420 425 430
 Asn Ser Val Gln Ala Phe Phe Asp Lys Ser Lys Phe Gly Glu Leu Pro

-continued

Leu Pro Arg Met Asp Leu Ser Glu Phe Phe Arg Pro Asn Asn Leu Thr
 850 855 860
 Met Phe Glu Asp Gly Lys Ser Ala Ser Ile Asp Leu Leu Leu Ile Leu
 865 870 875 880
 Asp Pro Leu Glu Glu Arg Thr Gln Met Ile Leu Ser Leu Val Glu Gln
 885 890 895
 Phe Arg Pro Leu Lys Phe Val Asn Ile Gln Val Ile Leu Met Pro Thr
 900 905 910
 Leu Glu Leu Asn Ile Val Pro Ile Arg Arg Ile Tyr Val Asp Asp Ala
 915 920 925
 Asp Ile Val Lys Ser Ile Thr Ser Glu Asp Ser Arg Ser Asp Pro Glu
 930 935 940
 Val Asp Ile Glu Met Asp Val Pro Asn Ser Phe Ile Val Asp Asn Asn
 945 950 955 960
 Tyr Arg Ile Lys Lys Leu Leu Ile Glu Leu His Ser Phe Ser Ser Lys
 965 970 975
 Thr Val Leu Ser Thr Gly Asn Ile Asp Gly Met Gly Gly Val Cys Leu
 980 985 990
 Ala Leu Val Asp Ser Ala Gly Asn Ile Ile Asp Lys Thr Thr Thr Met
 995 1000 1005
 Lys Thr Phe Gly Tyr Gly Gln Phe His Thr Asp Lys Phe Leu Lys
 1010 1015 1020
 Gly Cys Tyr Ile Lys Ser Cys Asp Ser Arg Tyr Thr Val Gln Ser
 1025 1030 1035
 Phe Ser Thr Asp Gly His Pro Asp Phe Ile Pro Ser Asp Ser Leu
 1040 1045 1050
 Asp Ile Leu Ser Tyr Asn Pro Gln Lys Ile Ala Val Lys Ile Ser
 1055 1060 1065
 Glu Glu Pro Thr His Glu Glu Glu Tyr Glu Glu Gly Arg Asn Asn
 1070 1075 1080
 Asp Thr Ile Ile Asn Ile Phe Thr Ile Ser Gly Pro Asp Glu Glu
 1085 1090 1095
 Glu Arg Tyr Met Gln Met Ile Leu Ser Ile Leu Ser Lys Cys Pro
 1100 1105 1110
 Glu Thr Gln Lys Val Asn Phe Phe Ile Leu Asp Gln Pro Phe Ile
 1115 1120 1125
 Ser Asp Thr Leu Arg Lys Ser Cys Glu Tyr Ile Asn Ser Ser Asp
 1130 1135 1140
 Glu Met Arg Gly Asn Val Ile Phe Leu Asn Tyr Glu Trp Pro Gln
 1145 1150 1155
 Trp Leu Arg Pro Gln Arg Phe Ser Ser Arg Arg Arg Asp Val Ser
 1160 1165 1170
 Arg Phe Leu Phe Leu Asp Val Leu Leu Pro Gln Asn Ile Ser Lys
 1175 1180 1185
 Val Leu Tyr Met Ser Pro Thr Glu Val Pro Leu Asp Pro Phe Asp
 1190 1195 1200
 Ile Phe Gln Phe Gln Gly Leu Lys Arg Ala Pro Leu Gly Leu Phe
 1205 1210 1215
 Arg Met Ser Gly Asp Gly Tyr Trp Lys Glu Gly Tyr Trp Glu Lys
 1220 1225 1230

-continued

Met Leu Arg Glu Asn Asn Leu Glu Phe Tyr Ser Thr Glu Pro Ala
 1235 1240 1245

Phe Leu Val Asn Leu Glu Arg Phe Arg Glu Leu Asp Ala Gly Asp
 1250 1255 1260

Lys Tyr Arg Ile His Tyr Gln Arg Ile Ser Thr Asp Ala Met Ser
 1265 1270 1275

Leu Val Asn Ile Gly Gln Asp Leu Val Asn Asn Leu Gln Leu Glu
 1280 1285 1290

Val Pro Ile Arg Phe Leu Lys Gly Ser Tyr Lys Lys Lys Leu Val
 1295 1300 1305

Ile Asn Asp Glu Cys Val Ser Glu Trp Lys Lys Lys Ile Asn Lys
 1310 1315 1320

Phe Ala Ser Ser Pro Gly Asp Glu Asp Val Pro Gly Glu Ser Val
 1325 1330 1335

Ser Ser Lys Tyr Gln Asp Ser Asp Asn Ala Ala Pro Leu His Asp
 1340 1345 1350

Glu Leu
 1355

<210> SEQ ID NO 5
 <211> LENGTH: 5616
 <212> TYPE: DNA
 <213> ORGANISM: Artificial sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Synthetic construct
 <220> FEATURE:
 <221> NAME/KEY: CDS
 <222> LOCATION: (1)..(5616)
 <223> OTHER INFORMATION: Saccharomyces cerevisiae s288c FKS1 (codon optimized)

<400> SEQUENCE: 5

atg aat acc gac cag caa ccg tac caa gga cag acc gac tat acc caa 48
 Met Asn Thr Asp Gln Gln Pro Tyr Gln Gly Gln Thr Asp Tyr Thr Gln
 1 5 10 15

ggc cca gga aac gga cag agc caa gag caa gac tac gat caa tac ggg 96
 Gly Pro Gly Asn Gly Gln Ser Gln Glu Gln Asp Tyr Asp Gln Tyr Gly
 20 25 30

cag ccg ctg tat ccg agt caa gcg gat ggc tac tac gac ccg aac gtt 144
 Gln Pro Leu Tyr Pro Ser Gln Ala Asp Gly Tyr Tyr Asp Pro Asn Val
 35 40 45

gcc gca ggc acg gaa gcc gac atg tat ggc cag cag ccc ccg aac gag 192
 Ala Ala Gly Thr Glu Ala Asp Met Tyr Gly Gln Gln Pro Pro Asn Glu
 50 55 60

tcg tat gac cag gat tat acc aac ggc gag tat tat ggc cag ccg ccc 240
 Ser Tyr Asp Gln Asp Tyr Thr Asn Gly Glu Tyr Tyr Gly Gln Pro Pro
 65 70 75 80

aac atg gcc gct caa gac ggc gag aat ttc agc gac ttc tcc tcg tat 288
 Asn Met Ala Ala Gln Asp Gly Glu Asn Phe Ser Asp Phe Ser Ser Tyr
 85 90 95

ggt ccg cct ggt acc ccg ggg tac gat tcc tat ggc ggg cag tac acg 336
 Gly Pro Pro Gly Thr Pro Gly Tyr Asp Ser Tyr Gly Gly Gln Tyr Thr
 100 105 110

gca tcg caa atg tcc tat ggc gag ccg aat agc tcg ggc acc agt acg 384
 Ala Ser Gln Met Ser Tyr Gly Glu Pro Asn Ser Ser Gly Thr Ser Thr
 115 120 125

ccg ata tac ggc aac tac gat ccg aac gcc atc gca atg gca ctg ccc 432
 Pro Ile Tyr Gly Asn Tyr Asp Pro Asn Ala Ile Ala Met Ala Leu Pro

-continued

130	135	140	
aac gag ccg tac ccc gcg tgg acg gcc gac tcg cag agc ccg gtc agc Asn Glu Pro Tyr Pro Ala Trp Thr Ala Asp Ser Gln Ser Pro Val Ser 145 150 155 160			480
atc gaa cag atc gag gac ata ttc atc gac ctc acc aat cgc ctc ggc Ile Glu Gln Ile Glu Asp Ile Phe Ile Asp Leu Thr Asn Arg Leu Gly 165 170 175			528
ttc cag cgc gac tcc atg cgc aac atg ttc gac cat ttc atg gtc ctc Phe Gln Arg Asp Ser Met Arg Asn Met Phe Asp His Phe Met Val Leu 180 185 190			576
ctc gac tcc cgc tcc tcg cgc atg agc ccg gac caa gcg ctc ctg tcc Leu Asp Ser Arg Ser Ser Arg Met Ser Pro Asp Gln Ala Leu Leu Ser 195 200 205			624
ttg cat gct gac tat att ggc ggc gac acc gcc aac tat aag aaa tgg Leu His Ala Asp Tyr Ile Gly Gly Asp Thr Ala Asn Tyr Lys Lys Trp 210 215 220			672
tat ttc gcc gcc cag ctc gac atg gac gac gag att ggc ttc cgg aat Tyr Phe Ala Ala Gln Leu Asp Met Asp Asp Glu Ile Gly Phe Arg Asn 225 230 235 240			720
atg tcc ctc ggg aag ctc tcg cgc aag gcc cgc aag gca aag aaa aag Met Ser Leu Gly Lys Leu Ser Arg Lys Ala Arg Lys Ala Lys Lys Lys 245 250 255			768
aac aag aag gca atg gaa gag gcg aat ccc gag gat acc gag gaa acc Asn Lys Lys Ala Met Glu Glu Ala Asn Pro Glu Asp Thr Glu Glu Thr 260 265 270			816
ctc aat aag att gag ggc gac aac agc ctt gag gct gcg gac ttc cgc Leu Asn Lys Ile Glu Gly Asp Asn Ser Leu Glu Ala Ala Asp Phe Arg 275 280 285			864
tgg aaa gcg aag atg aat cag ctg tcg ccc ctt gag cgc gtc cgc cac Trp Lys Ala Lys Met Asn Gln Leu Ser Pro Leu Glu Arg Val Arg His 290 295 300			912
atc gcg ctc tat ctc ctg tgt tgg ggg gaa gcc aat cag gtc cgc ttc Ile Ala Leu Tyr Leu Leu Cys Trp Gly Glu Ala Asn Gln Val Arg Phe 305 310 315 320			960
acc gcc gaa tgc ctc tgc ttc atc tac aag tgc gcg ctc gat tac ctg Thr Ala Glu Cys Leu Cys Phe Ile Tyr Lys Cys Ala Leu Asp Tyr Leu 325 330 335			1008
gac agc ccg ctc tgc cag cag cgg caa gaa ccc atg ccc gag ggt gac Asp Ser Pro Leu Cys Gln Gln Arg Gln Glu Pro Met Pro Glu Gly Asp 340 345 350			1056
ttc ctg aat cgc gtg atc acc cgg atc tat cac ttc ata cgc aac cag Phe Leu Asn Arg Val Ile Thr Pro Ile Tyr His Phe Ile Arg Asn Gln 355 360 365			1104
gtg tac gag att gtg gat ggc cgg ttc gtc aag cgc gag cgc gat cac Val Tyr Glu Ile Val Asp Gly Arg Phe Val Lys Arg Glu Arg Asp His 370 375 380			1152
aat aag atc gtg ggc tat gat gac ctg aac cag ctc ttt tgg tac ccg Asn Lys Ile Val Gly Tyr Asp Asp Leu Asn Gln Leu Phe Trp Tyr Pro 385 390 395 400			1200
gaa gga atc gcg aag ata gtt ctg gaa gat ggc acc aag ctt atc gag Glu Gly Ile Ala Lys Ile Val Leu Glu Asp Gly Thr Lys Leu Ile Glu 405 410 415			1248
ctc cgg ctt gag gag cgc tat ctg cgg ctg ggt gac gtc gtg tgg gac Leu Pro Leu Glu Glu Arg Tyr Leu Arg Leu Gly Asp Val Val Trp Asp 420 425 430			1296
gac gtg ttc ttc aaa acg tac aag gaa acg cgc acc tgg ctg cac ctt Asp Val Phe Phe Lys Thr Tyr Lys Glu Thr Arg Thr Trp Leu His Leu 1344			

-continued

435		440		445												
gtc	acg	aac	ttc	aat	cgc	atc	tgg	gtg	atg	cat	atc	tcg	att	ttc	tgg	1392
Val	Thr	Asn	Phe	Asn	Arg	Ile	Trp	Val	Met	His	Ile	Ser	Ile	Phe	Trp	
	450					455					460					
atg	tac	ttc	gca	tac	aac	tcg	ccg	acg	ttc	tat	acc	cac	aat	tat	cag	1440
Met	Tyr	Phe	Ala	Tyr	Asn	Ser	Pro	Thr	Phe	Tyr	Thr	His	Asn	Tyr	Gln	
	465				470					475					480	
caa	ctc	gtc	gac	aat	caa	ccg	ctg	gcc	gcg	tac	aag	tgg	gcg	tcg	tgc	1488
Gln	Leu	Val	Asp	Asn	Gln	Pro	Leu	Ala	Ala	Tyr	Lys	Trp	Ala	Ser	Cys	
				485					490					495		
gct	ctg	ggc	ggc	acc	gtg	gcg	tcc	ctc	atc	cag	att	gtc	gcc	acg	ctc	1536
Ala	Leu	Gly	Gly	Thr	Val	Ala	Ser	Leu	Ile	Gln	Ile	Val	Ala	Thr	Leu	
			500					505					510			
tgt	gag	tgg	tcc	ttc	gtc	ccg	ccg	aaa	tgg	gcg	gga	gcc	cag	cat	ctg	1584
Cys	Glu	Trp	Ser	Phe	Val	Pro	Arg	Lys	Trp	Ala	Gly	Ala	Gln	His	Leu	
		515					520					525				
tcg	cgc	ccg	ttc	tgg	ttc	ctg	tgc	atc	atc	ttc	ggg	atc	aac	ctg	ggc	1632
Ser	Arg	Arg	Phe	Trp	Phe	Leu	Cys	Ile	Ile	Phe	Gly	Ile	Asn	Leu	Gly	
	530					535					540					
ccg	atc	atc	ttc	gtg	ttc	gcc	tac	gac	aag	gac	acg	gtc	tat	tcc	acc	1680
Pro	Ile	Ile	Phe	Val	Phe	Ala	Tyr	Asp	Lys	Asp	Thr	Val	Tyr	Ser	Thr	
	545				550					555					560	
gca	gcg	cat	gtc	gtc	gca	gcg	gtc	atg	ttc	ttc	ggt	gcg	gtc	gcg	acc	1728
Ala	Ala	His	Val	Val	Ala	Ala	Val	Met	Phe	Phe	Val	Ala	Val	Ala	Thr	
				565					570					575		
atc	atc	ttt	ttc	tcc	atc	atg	cca	ctg	ggt	ggc	ctc	ttc	acc	agc	tat	1776
Ile	Ile	Phe	Phe	Ser	Ile	Met	Pro	Leu	Gly	Gly	Leu	Phe	Thr	Ser	Tyr	
			580					585					590			
atg	aag	aaa	tcg	act	cgc	ccg	tac	gtc	gct	agc	cag	acc	ttc	acg	gca	1824
Met	Lys	Lys	Ser	Thr	Arg	Arg	Tyr	Val	Ala	Ser	Gln	Thr	Phe	Thr	Ala	
		595					600					605				
gcg	ttc	gca	ccc	ctg	cat	ggc	ctc	gac	cgc	tgg	atg	agc	tac	ttg	gtg	1872
Ala	Phe	Ala	Pro	Leu	His	Gly	Leu	Asp	Arg	Trp	Met	Ser	Tyr	Leu	Val	
	610					615					620					
tgg	gtc	acg	gtg	ttc	gcg	gcc	aag	tat	tcc	gag	tcc	tac	tat	ttc	ctc	1920
Trp	Val	Thr	Val	Phe	Ala	Ala	Lys	Tyr	Ser	Glu	Ser	Tyr	Tyr	Phe	Leu	
	625				630					635					640	
gtg	ctg	tcc	ctc	cgc	gac	ccg	atc	cgc	atc	ctg	agc	acc	acc	gcc	atg	1968
Val	Leu	Ser	Leu	Arg	Asp	Pro	Ile	Arg	Ile	Leu	Ser	Thr	Thr	Ala	Met	
				645					650					655		
cgc	tgc	acc	ggg	gag	tac	tgg	tgg	ggt	gcg	gtg	ctc	tgc	aaa	gtc	cag	2016
Arg	Cys	Thr	Gly	Glu	Tyr	Trp	Trp	Gly	Ala	Val	Leu	Cys	Lys	Val	Gln	
			660					665					670			
ccc	aag	atc	ggt	ctt	ggc	ctg	gtg	atc	gcg	acg	gac	ttc	atc	ctc	ttt	2064
Pro	Lys	Ile	Val	Leu	Gly	Leu	Val	Ile	Ala	Thr	Asp	Phe	Ile	Leu	Phe	
		675				680							685			
ttc	ctt	gac	acc	tat	ctg	tgg	tac	att	atc	gtc	aac	acc	att	ttc	agc	2112
Phe	Leu	Asp	Thr	Tyr	Leu	Trp	Tyr	Ile	Ile	Val	Asn	Thr	Ile	Phe	Ser	
	690					695					700					
gtg	ggc	aag	tcg	ttc	tac	ctc	ggc	atc	agt	atc	ctg	acc	ccg	tgg	cgc	2160
Val	Gly	Lys	Ser	Phe	Tyr	Leu	Gly	Ile	Ser	Ile	Leu	Thr	Pro	Trp	Arg	
	705				710					715					720	
aac	atc	ttc	acc	ccg	ctc	ccc	aag	cgc	ata	tac	tcg	aag	att	ctg	gcc	2208
Asn	Ile	Phe	Thr	Arg	Leu	Pro	Lys	Arg	Ile	Tyr	Ser	Lys	Ile	Leu	Ala	
				725					730					735		
acc	act	gac	atg	gag	atc	aag	tat	aag	ccg	aag	gtc	ctc	att	agc	cag	2256
Thr	Thr	Asp	Met	Glu	Ile	Lys	Tyr	Lys	Pro	Lys	Val	Leu	Ile	Ser	Gln	

-continued

740			745			750										
gtg	tgg	aac	gcg	atc	ata	atc	tcg	atg	tat	cgg	gag	cac	ttg	ctg	gct	2304
Val	Trp	Asn	Ala	Ile	Ile	Ile	Ser	Met	Tyr	Arg	Glu	His	Leu	Leu	Ala	
		755					760					765				
atc	gac	cac	gtg	cag	aaa	ctg	ctg	tat	cat	caa	gtg	ccc	agt	gag	atc	2352
Ile	Asp	His	Val	Gln	Lys	Leu	Leu	Tyr	His	Gln	Val	Pro	Ser	Glu	Ile	
		770				775						780				
gag	ggg	aaa	cgg	acg	ctg	agg	gca	ccc	acc	ttc	ttt	gtg	agc	cag	gat	2400
Glu	Gly	Lys	Arg	Thr	Leu	Arg	Ala	Pro	Thr	Phe	Phe	Val	Ser	Gln	Asp	
					790							795			800	
gac	aat	aat	ttt	gaa	acc	gaa	ttc	ttc	cct	cgc	gat	tcc	gag	gcc	gag	2448
Asp	Asn	Asn	Phe	Glu	Thr	Glu	Phe	Phe	Pro	Arg	Asp	Ser	Glu	Ala	Glu	
				805						810					815	
cgg	cgc	atc	agc	ttc	ttt	gcc	caa	tcc	ctg	tcg	acg	ccc	atc	ccg	gag	2496
Arg	Arg	Ile	Ser	Phe	Phe	Ala	Gln	Ser	Leu	Ser	Thr	Pro	Ile	Pro	Glu	
				820					825						830	
ccc	ctg	ccg	gtg	gac	aac	atg	ccg	acc	ttt	acc	gtg	ctc	acg	ccc	cat	2544
Pro	Leu	Pro	Val	Asp	Asn	Met	Pro	Thr	Phe	Thr	Val	Leu	Thr	Pro	His	
				835					840						845	
tat	gcc	gag	cgc	atc	ctc	ctg	agc	ttg	cgc	gag	atc	atc	cgc	gag	gac	2592
Tyr	Ala	Glu	Arg	Ile	Leu	Leu	Ser	Leu	Arg	Glu	Ile	Ile	Arg	Glu	Asp	
				850				855							860	
gac	cag	ttc	tcg	cgg	gtt	acg	ctc	ctg	gag	tac	ctc	aag	caa	ctg	cat	2640
Asp	Gln	Phe	Ser	Arg	Val	Thr	Leu	Leu	Glu	Tyr	Leu	Lys	Gln	Leu	His	
						870									880	
ccg	gtg	gag	tgg	gag	tgc	ttc	gtc	aaa	gac	acc	aaa	atc	ctg	gcc	gag	2688
Pro	Val	Glu	Trp	Glu	Cys	Phe	Val	Lys	Asp	Thr	Lys	Ile	Leu	Ala	Glu	
					885										895	
gaa	act	gct	gcg	tac	gag	ggc	aat	gag	aac	gag	gcc	gag	aag	gaa	gat	2736
Glu	Thr	Ala	Ala	Tyr	Glu	Gly	Asn	Glu	Asn	Glu	Ala	Glu	Lys	Glu	Asp	
				900						905					910	
gca	ctg	aaa	tcc	cag	atc	gac	gat	ttg	cct	ttc	tat	tgt	atc	ggc	ttc	2784
Ala	Leu	Lys	Ser	Gln	Ile	Asp	Asp	Leu	Pro	Phe	Tyr	Cys	Ile	Gly	Phe	
				915					920						925	
aag	agt	gcg	gcc	ccg	gag	tac	acg	ctc	cgc	acc	cgg	atc	tgg	gcg	agt	2832
Lys	Ser	Ala	Ala	Pro	Glu	Tyr	Thr	Leu	Arg	Thr	Arg	Ile	Trp	Ala	Ser	
				930											940	
ctg	cgc	tcc	cag	acc	ctc	tac	cgc	acc	atc	tcg	ggc	ttc	atg	aac	tat	2880
Leu	Arg	Ser	Gln	Thr	Leu	Tyr	Arg	Thr	Ile	Ser	Gly	Phe	Met	Asn	Tyr	
					945										960	
tcg	cgc	gct	atc	aag	ctg	ctt	tat	cgg	gtc	gag	aac	ccc	gag	atc	gtg	2928
Ser	Arg	Ala	Ile	Lys	Leu	Leu	Tyr	Arg	Val	Glu	Asn	Pro	Glu	Ile	Val	
					965										975	
caa	atg	ttc	ggg	ggc	aat	gcc	gag	ggc	ctt	gag	cgc	gag	ctg	gag	aaa	2976
Gln	Met	Phe	Gly	Gly	Asn	Ala	Glu	Gly	Leu	Glu	Arg	Glu	Leu	Glu	Lys	
					980										990	
atg	gcg	agg	cgc	aag	ttc	aag	ttc	ctg	gtg	tcc	atg	cag	cgg	ctg	gcg	3024
Met	Ala	Arg	Arg	Lys	Phe	Lys	Phe	Leu	Val	Ser	Met	Gln	Arg	Leu	Ala	
				995											1005	
aag	ttt	aag	ttc	ctc	gaa	aat	gcc	gag	ttc	ctc	ctc	cgg	gcg	tac		3069
Lys	Phe	Lys	Phe	Leu	Glu	Asn	Ala	Glu	Phe	Leu	Leu	Arg	Ala	Tyr		
				1010											1020	
ccg	gac	ctc	cag	atc	gcc	tat	ctt	gac	gag	gaa	ccc	ccg	ctg	acg		3114
Pro	Asp	Leu	Gln	Ile	Ala	Tyr	Leu	Asp	Glu	Glu	Pro	Pro	Leu	Thr		
															1035	
gag	ggc	gag	gag	ccg	cgg	atc	tat	tcg	gcg	ctc	att	gac	ggc	cac		3159
Glu	Gly	Glu	Glu	Pro	Arg	Ile	Tyr	Ser	Ala	Leu	Ile	Asp	Gly	His		

-continued

1040	1045	1050	
tgc gag atc ctc gac aac ggt	cgg cgc agg cca aag	ttc cgc gtg	3204
Cys Glu Ile Leu Asp Asn Gly	Arg Arg Arg Pro Lys Phe Arg Val		
1055	1060	1065	
caa ctc agc ggc aat ccc att	ctg ggc gac ggc aaa	tcc gac aac	3249
Gln Leu Ser Gly Asn Pro Ile	Leu Gly Asp Gly Lys Ser Asp Asn		
1070	1075	1080	
caa aac cat gcc ctg atc ttc	tat agg ggt gag tat	att cag ctg	3294
Gln Asn His Ala Leu Ile Phe	Tyr Arg Gly Glu Tyr Ile Gln Leu		
1085	1090	1095	
atc gac gcg aac cag gac aat	tat ctt gag gaa tgc	ctc aag atc	3339
Ile Asp Ala Asn Gln Asp Asn	Tyr Leu Glu Glu Cys Leu Lys Ile		
1100	1105	1110	
cgc tcg gtc ctg gcc gag ttc	gag gag ctc aac gtc	gaa cag gtc	3384
Arg Ser Val Leu Ala Glu Phe	Glu Glu Leu Asn Val Glu Gln Val		
1115	1120	1125	
aac cct tat gct cgg ggc ctg	cgg tac gaa gaa cag	acc acg aac	3429
Asn Pro Tyr Ala Pro Gly Leu	Arg Tyr Glu Glu Gln Thr Thr Asn		
1130	1135	1140	
cat ccg gtc gcc atc gtc gga	gcg cgc gag tac att	ttc tcg gag	3474
His Pro Val Ala Ile Val Gly	Ala Arg Glu Tyr Ile Phe Ser Glu		
1145	1150	1155	
aat tcc ggc gtc ctc ggc gat	gtg gcg gca ggc aag	gag cag acc	3519
Asn Ser Gly Val Leu Gly Asp	Val Ala Ala Gly Lys Glu Gln Thr		
1160	1165	1170	
ttc ggc acc ctg ttc gcc cgc	acc ctc tcc cag att	ggg ggc aaa	3564
Phe Gly Thr Leu Phe Ala Arg	Thr Leu Ser Gln Ile Gly Gly Lys		
1175	1180	1185	
ctg cat tac ggc cat ccg gac	ttc ata aac gcg acc	ttc atg acc	3609
Leu His Tyr Gly His Pro Asp	Phe Ile Asn Ala Thr Phe Met Thr		
1190	1195	1200	
acc cgg ggt ggc gtc agc aag	gcc caa aag ggc ctc	cat ctt aac	3654
Thr Arg Gly Gly Val Ser Lys	Ala Gln Lys Gly Leu His Leu Asn		
1205	1210	1215	
gaa gat atc tac gcg ggt atg	aat gcc atg ctc agg	ggc ggt cgg	3699
Glu Asp Ile Tyr Ala Gly Met	Asn Ala Met Leu Arg Gly Gly Arg		
1220	1225	1230	
atc aag cat tgt gag tat tac	cag tgc gga aag ggc	agg gat ctg	3744
Ile Lys His Cys Glu Tyr Tyr	Gln Cys Gly Lys Gly Arg Asp Leu		
1235	1240	1245	
ggc ttc ggc acc atc ctc aat	ttc acc acc aag atc	ggg gca ggc	3789
Gly Phe Gly Thr Ile Leu Asn	Phe Thr Thr Lys Ile Gly Ala Gly		
1250	1255	1260	
atg gga gaa cag atg ttg agc	cgg gag tac tat tac	ctc ggg acg	3834
Met Gly Glu Gln Met Leu Ser	Arg Glu Tyr Tyr Tyr Leu Gly Thr		
1265	1270	1275	
cag ctc ccg gtc gat cgg ttc	ctg acc ttc tac tat	gcc cat ccg	3879
Gln Leu Pro Val Asp Arg Phe	Leu Thr Phe Tyr Tyr Ala His Pro		
1280	1285	1290	
ggg ttc cat ctg aat aac ctc	ttc atc caa ctg tcc	ctt cag atg	3924
Gly Phe His Leu Asn Asn Leu	Phe Ile Gln Leu Ser Leu Gln Met		
1295	1300	1305	
ttc atg ttg acg ctc gtg aac	ctg agt agc ctc gca	cat gag tcg	3969
Phe Met Leu Thr Leu Val Asn	Leu Ser Ser Leu Ala His Glu Ser		
1310	1315	1320	
atc atg tgc atc tac gat cgg	aat aag ccg aaa acc	gac gtc ctg	4014
Ile Met Cys Ile Tyr Asp Arg	Asn Lys Pro Lys Thr Asp Val Leu		

-continued

1325	1330	1335	
gtg ccg att ggc tgc tac aac ttc cag ccc gcg gtc gac tgg gtc			4059
Val Pro Ile Gly Cys Tyr Asn Phe Gln Pro Ala Val Asp Trp Val			
1340	1345	1350	
cgc cgc tat acg ctt agc atc ttt atc gtg ttc tgg atc gcg ttc			4104
Arg Arg Tyr Thr Leu Ser Ile Phe Ile Val Phe Trp Ile Ala Phe			
1355	1360	1365	
gtg ccg atc gtt gtg cag gag ctg atc gag cgc ggt ctg tgg aag			4149
Val Pro Ile Val Val Gln Glu Leu Ile Glu Arg Gly Leu Trp Lys			
1370	1375	1380	
gcc acg cag cgc ttc ttc tgc cat ctg ctg tcg ctg agt ccg atg			4194
Ala Thr Gln Arg Phe Phe Cys His Leu Leu Ser Leu Ser Pro Met			
1385	1390	1395	
ttc gag gtc ttt gcg ggc caa atc tat tcg agc gcg ctc ctg agc			4239
Phe Glu Val Phe Ala Gly Gln Ile Tyr Ser Ser Ala Leu Leu Ser			
1400	1405	1410	
gat ttg gcg atc ggg gga gcg cgc tac atc tcg acg ggt cgg ggc			4284
Asp Leu Ala Ile Gly Gly Ala Arg Tyr Ile Ser Thr Gly Arg Gly			
1415	1420	1425	
ttc gcc acc tcc cgc att cca ttc agc atc aag cgc ttc gcg ggc			4329
Phe Ala Thr Ser Arg Ile Pro Phe Ser Ile Lys Arg Phe Ala Gly			
1430	1435	1440	
tcg gcg atc tac atg ggc gca cgc tcg atg ttg atg ctg ctg ttc			4374
Ser Ala Ile Tyr Met Gly Ala Arg Ser Met Leu Met Leu Leu Phe			
1445	1450	1455	
ggc acc gtg gct cat tgg cag gcg ccg ctc ctg tgg ttc tgg gcg			4419
Gly Thr Val Ala His Trp Gln Ala Pro Leu Leu Trp Phe Trp Ala			
1460	1465	1470	
tcc ctg agc agc ctc atc ttc gcc ccc ttc gtg ttc aac ccg cat			4464
Ser Leu Ser Ser Leu Ile Phe Ala Pro Phe Val Phe Asn Pro His			
1475	1480	1485	
cag ttt gcg tgg gag gac ttt ttc ctg gac tac cgc gac tac atc			4509
Gln Phe Ala Trp Glu Asp Phe Phe Leu Asp Tyr Arg Asp Tyr Ile			
1490	1495	1500	
cgg tgg ctc tcc cgg gga aat aac cag tac cac cgc aat tcc tgg			4554
Arg Trp Leu Ser Arg Gly Asn Asn Gln Tyr His Arg Asn Ser Trp			
1505	1510	1515	
att ggc tat gtg cgg atg agc cgc gct cgc atc acc ggc ttc aag			4599
Ile Gly Tyr Val Arg Met Ser Arg Ala Arg Ile Thr Gly Phe Lys			
1520	1525	1530	
cgg aaa ctg gtc ggt gac gaa agc gag aaa gcc gcg ggc gac gcc			4644
Arg Lys Leu Val Gly Asp Glu Ser Glu Lys Ala Ala Gly Asp Ala			
1535	1540	1545	
tcg cgg gcc cac cgc acc aac ctg atc atg gcc gag atc atc ccc			4689
Ser Arg Ala His Arg Thr Asn Leu Ile Met Ala Glu Ile Ile Pro			
1550	1555	1560	
tgc gcc atc tat gcg gca ggg tgc ttc ata gcg ttc acc ttc atc			4734
Cys Ala Ile Tyr Ala Ala Gly Cys Phe Ile Ala Phe Thr Phe Ile			
1565	1570	1575	
aac gcc cag act ggc gtc aag acc acc gac gac gac cgg gtc aac			4779
Asn Ala Gln Thr Gly Val Lys Thr Thr Asp Asp Asp Arg Val Asn			
1580	1585	1590	
tcg gtg ctc cgc atc atc atc tgc acg ctg gcg ccg att gcg gtc			4824
Ser Val Leu Arg Ile Ile Ile Cys Thr Leu Ala Pro Ile Ala Val			
1595	1600	1605	
aat ctc ggg gtc ctc ttc ttc tgc atg ggc atg tcg tgc tgc tcc			4869
Asn Leu Gly Val Leu Phe Phe Cys Met Gly Met Ser Cys Cys Ser			

-continued

1610	1615	1620	
ggc cca ctg ttc ggc atg tgc tgc aag aaa acg ggc tgc gtc atg			4914
Gly Pro Leu Phe Gly Met Cys Cys Lys Lys Thr Gly Ser Val Met			
1625	1630	1635	
gcc ggc atc gcc cac ggc gtc gcg gtc att gtg cat atc gct ttc			4959
Ala Gly Ile Ala His Gly Val Ala Val Ile Val His Ile Ala Phe			
1640	1645	1650	
ttc atc gtg atg tgg gtt ctg gag tcc ttc aat ttc gtc cgg atg			5004
Phe Ile Val Met Trp Val Leu Glu Ser Phe Asn Phe Val Arg Met			
1655	1660	1665	
ctg atc ggc gtg gtg acg tgc atc cag tgc cag cgc ctc atc ttc			5049
Leu Ile Gly Val Val Thr Cys Ile Gln Cys Gln Arg Leu Ile Phe			
1670	1675	1680	
cac tgc atg acc gcc ctc atg ctc acg cgg gag ttc aaa aat gat			5094
His Cys Met Thr Ala Leu Met Leu Thr Arg Glu Phe Lys Asn Asp			
1685	1690	1695	
cat gcg aat acg gcc ttc tgg acc gcc aaa tgg tac ggc aag ggc			5139
His Ala Asn Thr Ala Phe Trp Thr Gly Lys Trp Tyr Gly Lys Gly			
1700	1705	1710	
atg ggc tat atg gcc tgg acg caa ccc tcg cgc gag ctg acg gcc			5184
Met Gly Tyr Met Ala Trp Thr Gln Pro Ser Arg Glu Leu Thr Ala			
1715	1720	1725	
aag gtc atc gag ttg tcc gag ttt gcc gct gac ttc gtc ctc gcc			5229
Lys Val Ile Glu Leu Ser Glu Phe Ala Ala Asp Phe Val Leu Gly			
1730	1735	1740	
cat gtt atc ttg atc tgc cag ctg ccg ctg ata atc ata ccg aag			5274
His Val Ile Leu Ile Cys Gln Leu Pro Leu Ile Ile Ile Pro Lys			
1745	1750	1755	
atc gac aag ttt cat tcc atc atg ctg ttc tgg ctg aaa ccg tcg			5319
Ile Asp Lys Phe His Ser Ile Met Leu Phe Trp Leu Lys Pro Ser			
1760	1765	1770	
cgc cag att agg ccc cct atc tac tcg ctc aaa cag act agg ctc			5364
Arg Gln Ile Arg Pro Pro Ile Tyr Ser Leu Lys Gln Thr Arg Leu			
1775	1780	1785	
cgg aaa cgc atg gtc aag aaa tac tgc tcg ctg tat ttc ctc gtg			5409
Arg Lys Arg Met Val Lys Lys Tyr Cys Ser Leu Tyr Phe Leu Val			
1790	1795	1800	
ctg gcc ata ttc gcg ggc tgc atc atc gga ccg gcc gtg gcg agc			5454
Leu Ala Ile Phe Ala Gly Cys Ile Ile Gly Pro Ala Val Ala Ser			
1805	1810	1815	
gcc aag atc cat aag cat atc ggg gat tcc ctg gac ggt gtc gtc			5499
Ala Lys Ile His Lys His Ile Gly Asp Ser Leu Asp Gly Val Val			
1820	1825	1830	
cac aac ctg ttc cag ccg atc aac act acc aac aac gac acc ggc			5544
His Asn Leu Phe Gln Pro Ile Asn Thr Thr Asn Asn Asp Thr Gly			
1835	1840	1845	
tcc cag atg tcg acc tac cag tcg cac tac tac acc cac acc ccg			5589
Ser Gln Met Ser Thr Tyr Gln Ser His Tyr Tyr Thr His Thr Pro			
1850	1855	1860	
tcc ctt aag acc tgg agc acc ata aag			5616
Ser Leu Lys Thr Trp Ser Thr Ile Lys			
1865	1870		

<210> SEQ ID NO 6

<211> LENGTH: 1872

<212> TYPE: PRT

<213> ORGANISM: Artificial sequence

<220> FEATURE:

-continued

<223> OTHER INFORMATION: Synthetic Construct

<400> SEQUENCE: 6

```

Met Asn Thr Asp Gln Gln Pro Tyr Gln Gly Gln Thr Asp Tyr Thr Gln
1      5      10      15
Gly Pro Gly Asn Gly Gln Ser Gln Glu Gln Asp Tyr Asp Gln Tyr Gly
20      25      30
Gln Pro Leu Tyr Pro Ser Gln Ala Asp Gly Tyr Tyr Asp Pro Asn Val
35      40      45
Ala Ala Gly Thr Glu Ala Asp Met Tyr Gly Gln Gln Pro Pro Asn Glu
50      55      60
Ser Tyr Asp Gln Asp Tyr Thr Asn Gly Glu Tyr Tyr Gly Gln Pro Pro
65      70      75      80
Asn Met Ala Ala Gln Asp Gly Glu Asn Phe Ser Asp Phe Ser Ser Tyr
85      90      95
Gly Pro Pro Gly Thr Pro Gly Tyr Asp Ser Tyr Gly Gly Gln Tyr Thr
100     105     110
Ala Ser Gln Met Ser Tyr Gly Glu Pro Asn Ser Ser Gly Thr Ser Thr
115     120     125
Pro Ile Tyr Gly Asn Tyr Asp Pro Asn Ala Ile Ala Met Ala Leu Pro
130     135     140
Asn Glu Pro Tyr Pro Ala Trp Thr Ala Asp Ser Gln Ser Pro Val Ser
145     150     155     160
Ile Glu Gln Ile Glu Asp Ile Phe Ile Asp Leu Thr Asn Arg Leu Gly
165     170     175
Phe Gln Arg Asp Ser Met Arg Asn Met Phe Asp His Phe Met Val Leu
180     185     190
Leu Asp Ser Arg Ser Ser Arg Met Ser Pro Asp Gln Ala Leu Leu Ser
195     200     205
Leu His Ala Asp Tyr Ile Gly Gly Asp Thr Ala Asn Tyr Lys Lys Trp
210     215     220
Tyr Phe Ala Ala Gln Leu Asp Met Asp Asp Glu Ile Gly Phe Arg Asn
225     230     235     240
Met Ser Leu Gly Lys Leu Ser Arg Lys Ala Arg Lys Ala Lys Lys Lys
245     250     255
Asn Lys Lys Ala Met Glu Glu Ala Asn Pro Glu Asp Thr Glu Glu Thr
260     265     270
Leu Asn Lys Ile Glu Gly Asp Asn Ser Leu Glu Ala Ala Asp Phe Arg
275     280     285
Trp Lys Ala Lys Met Asn Gln Leu Ser Pro Leu Glu Arg Val Arg His
290     295     300
Ile Ala Leu Tyr Leu Leu Cys Trp Gly Glu Ala Asn Gln Val Arg Phe
305     310     315     320
Thr Ala Glu Cys Leu Cys Phe Ile Tyr Lys Cys Ala Leu Asp Tyr Leu
325     330     335
Asp Ser Pro Leu Cys Gln Gln Arg Gln Glu Pro Met Pro Glu Gly Asp
340     345     350
Phe Leu Asn Arg Val Ile Thr Pro Ile Tyr His Phe Ile Arg Asn Gln
355     360     365
Val Tyr Glu Ile Val Asp Gly Arg Phe Val Lys Arg Glu Arg Asp His
370     375     380

```

-continued

Asn	Lys	Ile	Val	Gly	Tyr	Asp	Asp	Leu	Asn	Gln	Leu	Phe	Trp	Tyr	Pro
385					390					395					400
Glu	Gly	Ile	Ala	Lys	Ile	Val	Leu	Glu	Asp	Gly	Thr	Lys	Leu	Ile	Glu
				405					410					415	
Leu	Pro	Leu	Glu	Glu	Arg	Tyr	Leu	Arg	Leu	Gly	Asp	Val	Val	Trp	Asp
			420					425					430		
Asp	Val	Phe	Phe	Lys	Thr	Tyr	Lys	Glu	Thr	Arg	Thr	Trp	Leu	His	Leu
		435					440					445			
Val	Thr	Asn	Phe	Asn	Arg	Ile	Trp	Val	Met	His	Ile	Ser	Ile	Phe	Trp
	450					455					460				
Met	Tyr	Phe	Ala	Tyr	Asn	Ser	Pro	Thr	Phe	Tyr	Thr	His	Asn	Tyr	Gln
465					470					475					480
Gln	Leu	Val	Asp	Asn	Gln	Pro	Leu	Ala	Ala	Tyr	Lys	Trp	Ala	Ser	Cys
				485					490						495
Ala	Leu	Gly	Gly	Thr	Val	Ala	Ser	Leu	Ile	Gln	Ile	Val	Ala	Thr	Leu
			500					505					510		
Cys	Glu	Trp	Ser	Phe	Val	Pro	Arg	Lys	Trp	Ala	Gly	Ala	Gln	His	Leu
		515					520					525			
Ser	Arg	Arg	Phe	Trp	Phe	Leu	Cys	Ile	Ile	Phe	Gly	Ile	Asn	Leu	Gly
	530					535					540				
Pro	Ile	Ile	Phe	Val	Phe	Ala	Tyr	Asp	Lys	Asp	Thr	Val	Tyr	Ser	Thr
545					550					555					560
Ala	Ala	His	Val	Val	Ala	Ala	Val	Met	Phe	Phe	Val	Ala	Val	Ala	Thr
				565					570						575
Ile	Ile	Phe	Phe	Ser	Ile	Met	Pro	Leu	Gly	Gly	Leu	Phe	Thr	Ser	Tyr
			580					585					590		
Met	Lys	Lys	Ser	Thr	Arg	Arg	Tyr	Val	Ala	Ser	Gln	Thr	Phe	Thr	Ala
		595					600					605			
Ala	Phe	Ala	Pro	Leu	His	Gly	Leu	Asp	Arg	Trp	Met	Ser	Tyr	Leu	Val
	610					615					620				
Trp	Val	Thr	Val	Phe	Ala	Ala	Lys	Tyr	Ser	Glu	Ser	Tyr	Tyr	Phe	Leu
625					630					635					640
Val	Leu	Ser	Leu	Arg	Asp	Pro	Ile	Arg	Ile	Leu	Ser	Thr	Thr	Ala	Met
				645					650						655
Arg	Cys	Thr	Gly	Glu	Tyr	Trp	Trp	Gly	Ala	Val	Leu	Cys	Lys	Val	Gln
			660					665					670		
Pro	Lys	Ile	Val	Leu	Gly	Leu	Val	Ile	Ala	Thr	Asp	Phe	Ile	Leu	Phe
		675					680					685			
Phe	Leu	Asp	Thr	Tyr	Leu	Trp	Tyr	Ile	Ile	Val	Asn	Thr	Ile	Phe	Ser
	690					695					700				
Val	Gly	Lys	Ser	Phe	Tyr	Leu	Gly	Ile	Ser	Ile	Leu	Thr	Pro	Trp	Arg
705					710					715					720
Asn	Ile	Phe	Thr	Arg	Leu	Pro	Lys	Arg	Ile	Tyr	Ser	Lys	Ile	Leu	Ala
				725					730						735
Thr	Thr	Asp	Met	Glu	Ile	Lys	Tyr	Lys	Pro	Lys	Val	Leu	Ile	Ser	Gln
			740					745					750		
Val	Trp	Asn	Ala	Ile	Ile	Ile	Ser	Met	Tyr	Arg	Glu	His	Leu	Leu	Ala
		755						760				765			
Ile	Asp	His	Val	Gln	Lys	Leu	Leu	Tyr	His	Gln	Val	Pro	Ser	Glu	Ile
	770					775					780				
Glu	Gly	Lys	Arg	Thr	Leu	Arg	Ala	Pro	Thr	Phe	Phe	Val	Ser	Gln	Asp

-continued

785	790	795	800
Asp Asn Asn Phe Glu Thr	Glu Phe Phe Pro Arg Asp Ser Glu Ala Glu		
	805	810	815
Arg Arg Ile Ser Phe Phe Ala Gln Ser Leu Ser Thr Pro Ile Pro Glu		825	830
	820		
Pro Leu Pro Val Asp Asn Met Pro Thr Phe Thr Val Leu Thr Pro His		840	845
	835		
Tyr Ala Glu Arg Ile Leu Leu Ser Leu Arg Glu Ile Ile Arg Glu Asp		855	860
	850		
Asp Gln Phe Ser Arg Val Thr Leu Leu Glu Tyr Leu Lys Gln Leu His		870	875
	865		880
Pro Val Glu Trp Glu Cys Phe Val Lys Asp Thr Lys Ile Leu Ala Glu		890	895
	885		
Glu Thr Ala Ala Tyr Glu Gly Asn Glu Asn Glu Ala Glu Lys Glu Asp		905	910
	900		
Ala Leu Lys Ser Gln Ile Asp Asp Leu Pro Phe Tyr Cys Ile Gly Phe		920	925
	915		
Lys Ser Ala Ala Pro Glu Tyr Thr Leu Arg Thr Arg Ile Trp Ala Ser		935	940
	930		
Leu Arg Ser Gln Thr Leu Tyr Arg Thr Ile Ser Gly Phe Met Asn Tyr		950	955
	945		960
Ser Arg Ala Ile Lys Leu Leu Tyr Arg Val Glu Asn Pro Glu Ile Val		970	975
	965		
Gln Met Phe Gly Gly Asn Ala Glu Gly Leu Glu Arg Glu Leu Glu Lys		985	990
	980		
Met Ala Arg Arg Lys Phe Lys Phe Leu Val Ser Met Gln Arg Leu Ala		1000	1005
	995		
Lys Phe Lys Phe Leu Glu Asn Ala Glu Phe Leu Leu Arg Ala Tyr		1015	1020
	1010		
Pro Asp Leu Gln Ile Ala Tyr Leu Asp Glu Glu Pro Pro Leu Thr		1030	1035
	1025		
Glu Gly Glu Glu Pro Arg Ile Tyr Ser Ala Leu Ile Asp Gly His		1045	1050
	1040		
Cys Glu Ile Leu Asp Asn Gly Arg Arg Arg Pro Lys Phe Arg Val		1060	1065
	1055		
Gln Leu Ser Gly Asn Pro Ile Leu Gly Asp Gly Lys Ser Asp Asn		1075	1080
	1070		
Gln Asn His Ala Leu Ile Phe Tyr Arg Gly Glu Tyr Ile Gln Leu		1090	1095
	1085		
Ile Asp Ala Asn Gln Asp Asn Tyr Leu Glu Glu Cys Leu Lys Ile		1105	1110
	1100		
Arg Ser Val Leu Ala Glu Phe Glu Glu Leu Asn Val Glu Gln Val		1120	1125
	1115		
Asn Pro Tyr Ala Pro Gly Leu Arg Tyr Glu Glu Gln Thr Thr Asn		1135	1140
	1130		
His Pro Val Ala Ile Val Gly Ala Arg Glu Tyr Ile Phe Ser Glu		1150	1155
	1145		
Asn Ser Gly Val Leu Gly Asp Val Ala Ala Gly Lys Glu Gln Thr		1165	1170
	1160		
Phe Gly Thr Leu Phe Ala Arg Thr Leu Ser Gln Ile Gly Gly Lys		1180	1185
	1175		

-continued

Leu	His	Tyr	Gly	His	Pro	Asp	Phe	Ile	Asn	Ala	Thr	Phe	Met	Thr
1190						1195					1200			
Thr	Arg	Gly	Gly	Val	Ser	Lys	Ala	Gln	Lys	Gly	Leu	His	Leu	Asn
1205						1210					1215			
Glu	Asp	Ile	Tyr	Ala	Gly	Met	Asn	Ala	Met	Leu	Arg	Gly	Gly	Arg
1220						1225					1230			
Ile	Lys	His	Cys	Glu	Tyr	Tyr	Gln	Cys	Gly	Lys	Gly	Arg	Asp	Leu
1235						1240					1245			
Gly	Phe	Gly	Thr	Ile	Leu	Asn	Phe	Thr	Thr	Lys	Ile	Gly	Ala	Gly
1250						1255					1260			
Met	Gly	Glu	Gln	Met	Leu	Ser	Arg	Glu	Tyr	Tyr	Tyr	Leu	Gly	Thr
1265						1270					1275			
Gln	Leu	Pro	Val	Asp	Arg	Phe	Leu	Thr	Phe	Tyr	Tyr	Ala	His	Pro
1280						1285					1290			
Gly	Phe	His	Leu	Asn	Asn	Leu	Phe	Ile	Gln	Leu	Ser	Leu	Gln	Met
1295						1300					1305			
Phe	Met	Leu	Thr	Leu	Val	Asn	Leu	Ser	Ser	Leu	Ala	His	Glu	Ser
1310						1315					1320			
Ile	Met	Cys	Ile	Tyr	Asp	Arg	Asn	Lys	Pro	Lys	Thr	Asp	Val	Leu
1325						1330					1335			
Val	Pro	Ile	Gly	Cys	Tyr	Asn	Phe	Gln	Pro	Ala	Val	Asp	Trp	Val
1340						1345					1350			
Arg	Arg	Tyr	Thr	Leu	Ser	Ile	Phe	Ile	Val	Phe	Trp	Ile	Ala	Phe
1355						1360					1365			
Val	Pro	Ile	Val	Val	Gln	Glu	Leu	Ile	Glu	Arg	Gly	Leu	Trp	Lys
1370						1375					1380			
Ala	Thr	Gln	Arg	Phe	Phe	Cys	His	Leu	Leu	Ser	Leu	Ser	Pro	Met
1385						1390					1395			
Phe	Glu	Val	Phe	Ala	Gly	Gln	Ile	Tyr	Ser	Ser	Ala	Leu	Leu	Ser
1400						1405					1410			
Asp	Leu	Ala	Ile	Gly	Gly	Ala	Arg	Tyr	Ile	Ser	Thr	Gly	Arg	Gly
1415						1420					1425			
Phe	Ala	Thr	Ser	Arg	Ile	Pro	Phe	Ser	Ile	Lys	Arg	Phe	Ala	Gly
1430						1435					1440			
Ser	Ala	Ile	Tyr	Met	Gly	Ala	Arg	Ser	Met	Leu	Met	Leu	Leu	Phe
1445						1450					1455			
Gly	Thr	Val	Ala	His	Trp	Gln	Ala	Pro	Leu	Leu	Trp	Phe	Trp	Ala
1460						1465					1470			
Ser	Leu	Ser	Ser	Leu	Ile	Phe	Ala	Pro	Phe	Val	Phe	Asn	Pro	His
1475						1480					1485			
Gln	Phe	Ala	Trp	Glu	Asp	Phe	Phe	Leu	Asp	Tyr	Arg	Asp	Tyr	Ile
1490						1495					1500			
Arg	Trp	Leu	Ser	Arg	Gly	Asn	Asn	Gln	Tyr	His	Arg	Asn	Ser	Trp
1505						1510					1515			
Ile	Gly	Tyr	Val	Arg	Met	Ser	Arg	Ala	Arg	Ile	Thr	Gly	Phe	Lys
1520						1525					1530			
Arg	Lys	Leu	Val	Gly	Asp	Glu	Ser	Glu	Lys	Ala	Ala	Gly	Asp	Ala
1535						1540					1545			
Ser	Arg	Ala	His	Arg	Thr	Asn	Leu	Ile	Met	Ala	Glu	Ile	Ile	Pro
1550						1555					1560			

-continued

Cys	Ala	Ile	Tyr	Ala	Ala	Gly	Cys	Phe	Ile	Ala	Phe	Thr	Phe	Ile
1565						1570					1575			
Asn	Ala	Gln	Thr	Gly	Val	Lys	Thr	Thr	Asp	Asp	Asp	Arg	Val	Asn
1580						1585					1590			
Ser	Val	Leu	Arg	Ile	Ile	Ile	Cys	Thr	Leu	Ala	Pro	Ile	Ala	Val
1595						1600					1605			
Asn	Leu	Gly	Val	Leu	Phe	Phe	Cys	Met	Gly	Met	Ser	Cys	Cys	Ser
1610						1615					1620			
Gly	Pro	Leu	Phe	Gly	Met	Cys	Cys	Lys	Lys	Thr	Gly	Ser	Val	Met
1625						1630					1635			
Ala	Gly	Ile	Ala	His	Gly	Val	Ala	Val	Ile	Val	His	Ile	Ala	Phe
1640						1645					1650			
Phe	Ile	Val	Met	Trp	Val	Leu	Glu	Ser	Phe	Asn	Phe	Val	Arg	Met
1655						1660					1665			
Leu	Ile	Gly	Val	Val	Thr	Cys	Ile	Gln	Cys	Gln	Arg	Leu	Ile	Phe
1670						1675					1680			
His	Cys	Met	Thr	Ala	Leu	Met	Leu	Thr	Arg	Glu	Phe	Lys	Asn	Asp
1685						1690					1695			
His	Ala	Asn	Thr	Ala	Phe	Trp	Thr	Gly	Lys	Trp	Tyr	Gly	Lys	Gly
1700						1705					1710			
Met	Gly	Tyr	Met	Ala	Trp	Thr	Gln	Pro	Ser	Arg	Glu	Leu	Thr	Ala
1715						1720					1725			
Lys	Val	Ile	Glu	Leu	Ser	Glu	Phe	Ala	Ala	Asp	Phe	Val	Leu	Gly
1730						1735					1740			
His	Val	Ile	Leu	Ile	Cys	Gln	Leu	Pro	Leu	Ile	Ile	Ile	Pro	Lys
1745						1750					1755			
Ile	Asp	Lys	Phe	His	Ser	Ile	Met	Leu	Phe	Trp	Leu	Lys	Pro	Ser
1760						1765					1770			
Arg	Gln	Ile	Arg	Pro	Pro	Ile	Tyr	Ser	Leu	Lys	Gln	Thr	Arg	Leu
1775						1780					1785			
Arg	Lys	Arg	Met	Val	Lys	Lys	Tyr	Cys	Ser	Leu	Tyr	Phe	Leu	Val
1790						1795					1800			
Leu	Ala	Ile	Phe	Ala	Gly	Cys	Ile	Ile	Gly	Pro	Ala	Val	Ala	Ser
1805						1810					1815			
Ala	Lys	Ile	His	Lys	His	Ile	Gly	Asp	Ser	Leu	Asp	Gly	Val	Val
1820						1825					1830			
His	Asn	Leu	Phe	Gln	Pro	Ile	Asn	Thr	Thr	Asn	Asn	Asp	Thr	Gly
1835						1840					1845			
Ser	Gln	Met	Ser	Thr	Tyr	Gln	Ser	His	Tyr	Tyr	Thr	His	Thr	Pro
1850						1855					1860			
Ser	Leu	Lys	Thr	Trp	Ser	Thr	Ile	Lys						
1865						1870								

<210> SEQ ID NO 7

<211> LENGTH: 5661

<212> TYPE: DNA

<213> ORGANISM: Artificial sequence

<220> FEATURE:

<223> OTHER INFORMATION: Synthetic construct

<220> FEATURE:

<221> NAME/KEY: CDS

<222> LOCATION: (1)..(5661)

<223> OTHER INFORMATION: Saccharomyces cerevisiae FKS2 (codon optimized)

<400> SEQUENCE: 7

-continued

atg tcc tac aac gat ccc aat ctc aat ggc cag tat tac tcc aac ggc	48
Met Ser Tyr Asn Asp Pro Asn Leu Asn Gly Gln Tyr Tyr Ser Asn Gly	
1 5 10 15	
gac ggc acc ggc gat ggc aat tat ccg acc tac cag gtg acc cag gat	96
Asp Gly Thr 20 Asp Gly Asn Tyr 25 Thr Tyr Gln Val Thr 30 Gln Asp	
caa agc gcg tac gac gag tac ggc cag ccc atc tat acc caa aat caa	144
Gln Ser Ala Tyr Asp Glu Tyr Gly Gln Pro Ile Tyr Thr Gln Asn Gln	
35 40 45	
ctg gac gat ggc tac tat gac ccg aat gaa caa tac gtg gat ggc acc	192
Leu Asp Asp Gly Tyr Tyr Asp Pro Asn Glu Gln Tyr Val Asp Gly Thr	
50 55 60	
cag ttc ccg cag ggc cag gac ccg tcg cag gac caa ggc ccg tat aac	240
Gln Phe Pro Gln Gly Gln Asp Pro Ser Gln Asp Gln Gly Pro Tyr Asn	
65 70 75 80	
aat gac gcc agc tat tac aac cag ccg ccg aac atg atg aac ccg tcg	288
Asn Asp Ala Ser Tyr Tyr Asn Gln Pro Pro Asn Met Met Asn Pro Ser	
85 90 95	
tcc cag gac ggt gag aat ttc tcg gac ttc agc tcg tat gga ccg ccc	336
Ser Gln Asp Gly Glu Asn Phe Ser Asp Phe Ser Ser Tyr Gly Pro Pro	
100 105 110	
tcg gga acg tac cct aat gac cag tat acg ccc agt cag atg tcc tac	384
Ser Gly Thr Tyr Pro Asn Asp Gln Tyr Thr Pro Ser Gln Met Ser Tyr	
115 120 125	
ccg gat cag gac ggc tcg tcg ggt gcg tcg acg ccg tat ggc aac ggc	432
Pro Asp Gln Asp Gly Ser Ser Gly Ala Ser Thr Pro Tyr Gly Asn Gly	
130 135 140	
gtc gtc aat ggg aac ggc cag tat tat gat ccg aat gcg atc gag atg	480
Val Val Asn Gly Asn Gly Gln Tyr Tyr Asp Pro Asn Ala Ile Glu Met	
145 150 155 160	
gcg ctg ccc aat gac ccg tat ccg gcg tgg acc gcc gat ccc cag tcc	528
Ala Leu Pro Asn Asp Pro Tyr Pro Ala Trp Thr Ala Asp Pro Gln Ser	
165 170 175	
cct ctc ccc atc gag caa att gag gac atc ttc ata gat ctc acc aat	576
Pro Leu Pro Ile Glu Gln Ile Glu Asp Ile Phe Ile Asp Leu Thr Asn	
180 185 190	
aag ttc ggc ttc cag cgc gac tcg atg ccg aac atg ttc gac cac ttc	624
Lys Phe Gly Phe Gln Arg Asp Ser Met Arg Asn Met Phe Asp His Phe	
195 200 205	
atg acc ctg ctg gat tcg cgc agc tcg cgc atg tcg ccg gaa cag gcc	672
Met Thr Leu Leu Asp Ser Arg Ser Ser Arg Met Ser Pro Glu Gln Ala	
210 215 220	
ttg ctc tcc ctc cat gcc gac tac att ggc ggc gac acc gcc aac tac	720
Leu Leu Ser Leu His Ala Asp Tyr Ile Gly Gly Asp Thr Ala Asn Tyr	
225 230 235 240	
aag aaa tgg tat ttt gcg gcc caa ctg gat atg gac gat gag atc ggc	768
Lys Lys Trp Tyr Phe Ala Ala Gln Leu Asp Met Asp Asp Glu Ile Gly	
245 250 255	
ttc ccg aac atg aaa ctc ggc aag ctg tcc cgc aaa gcg cgc aaa gcg	816
Phe Arg Asn Met Lys Leu Gly Lys Leu Ser Arg Lys Ala Arg Lys Ala	
260 265 270	
aag aag aaa aac aaa aag gcc atg caa gag gat gag gac act gaa gag	864
Lys Lys Lys Asn Lys Lys Ala Met Gln Glu Asp Glu Asp Thr Glu Glu	
275 280 285	
act ctg aac cag atc gag ggc gat aac tcc ctc gaa gcc gct gac ttc	912
Thr Leu Asn Gln Ile Glu Gly Asp Asn Ser Leu Glu Ala Ala Asp Phe	
290 295 300	

-continued

cgg tgg aag agt aaa atg aat caa ctc agc ccc ttc gag atg gtg cgg	960
Arg Trp Lys Ser Lys Met Asn Gln Leu Ser Pro Phe Glu Met Val Arg	
305 310 315 320	
caa atc gcc ctc ttc ctc ctg tgc tgg ggc gag gcc aat caa gtg cgg	1008
Gln Ile Ala Leu Phe Leu Leu Cys Trp Gly Glu Ala Asn Gln Val Arg	
325 330 335	
ttc acc ccg gag tgc ctg tgc ttt atc tac aag tgc gcg agc gac tac	1056
Phe Thr Pro Glu Cys Leu Cys Phe Ile Tyr Lys Cys Ala Ser Asp Tyr	
340 345 350	
ctg gac tcc gcc caa tgc cag cag cgg ccc gac ccc ctg ccc gag ggc	1104
Leu Asp Ser Ala Gln Cys Gln Gln Arg Pro Asp Pro Leu Pro Glu Gly	
355 360 365	
gac ttt ctc aat ccg gtg atc acg ccc ttg tac cgc ttc att cgc agc	1152
Asp Phe Leu Asn Arg Val Ile Thr Pro Leu Tyr Arg Phe Ile Arg Ser	
370 375 380	
cag gtg tac gag atc gtc gac ggc cgc tac gtg aag tcc gag aag gac	1200
Gln Val Tyr Glu Ile Val Asp Gly Arg Tyr Val Lys Ser Glu Lys Asp	
385 390 395 400	
cat aac aag gtc att ggt tat gac gac gtc aac caa ctt ttc tgg tat	1248
His Asn Lys Val Ile Gly Tyr Asp Asp Val Asn Gln Leu Phe Trp Tyr	
405 410 415	
ccg gaa ggc atc gcc aag atc gtg atg gag gac ggc acg cgc ctt atc	1296
Pro Glu Gly Ile Ala Lys Ile Val Met Glu Asp Gly Thr Arg Leu Ile	
420 425 430	
gac ctc ccc gct gag gaa cgg tac ctc aag ctc ggc gag atc ccc tgg	1344
Asp Leu Pro Ala Glu Glu Arg Tyr Leu Lys Leu Gly Glu Ile Pro Trp	
435 440 445	
gac gat gtg ttc ttc aag acc tat aag gaa acc cgc tcc tgg ctc cat	1392
Asp Asp Val Phe Phe Lys Thr Tyr Lys Glu Thr Arg Ser Trp Leu His	
450 455 460	
ctg gtg acc aac ttc aat cgg att tgg att atg cac gtg tat tgg atg	1440
Leu Val Thr Asn Phe Asn Arg Ile Trp Ile Met His Val Tyr Trp Met	
465 470 475 480	
tat tgt gca tac aat gcc ccc acc ttc tat acg cac aat tac caa cag	1488
Tyr Cys Ala Tyr Asn Ala Pro Thr Phe Tyr Thr His Asn Tyr Gln Gln	
485 490 495	
ctt gtc gac aac cag ccg ctc gcc gca tac aag tgg gcc acc gct gcc	1536
Leu Val Asp Asn Gln Pro Leu Ala Ala Tyr Lys Trp Ala Thr Ala Ala	
500 505 510	
ctc ggg ggg acg gtc gcg tcg ttg atc cag gtc gcg gcg acg ctg tgc	1584
Leu Gly Gly Thr Val Ala Ser Leu Ile Gln Val Ala Ala Thr Leu Cys	
515 520 525	
gag tgg tcc ttc gtg ccg cgg aaa tgg gca ggc gcg cag cac ctg tcc	1632
Glu Trp Ser Phe Val Pro Arg Lys Trp Ala Gly Ala Gln His Leu Ser	
530 535 540	
cgg cgc ttc tgg ttc ctg tgc gtc atc atg ggc atc aat ctt ggc ccc	1680
Arg Arg Phe Trp Phe Leu Cys Val Ile Met Gly Ile Asn Leu Gly Pro	
545 550 555 560	
gtc atc ttc gtg ttc gcc tat gac aaa gat acg gtc tac tcg acg gcc	1728
Val Ile Phe Val Phe Ala Tyr Asp Lys Asp Thr Val Tyr Ser Thr Ala	
565 570 575	
gca cac gtc gtg ggc gca gtt atg ttc ttt gtc gcc gtg gcg act ctg	1776
Ala His Val Val Gly Ala Val Met Phe Phe Val Ala Val Ala Thr Leu	
580 585 590	
gtc ttt ttc agc gtc atg ccg ctg ggt ggc ctg ttc acc agt tat atg	1824
Val Phe Phe Ser Val Met Pro Leu Gly Gly Leu Phe Thr Ser Tyr Met	
595 600 605	

-continued

aag aaa agt acc cgg tcg tac gtc gcg agt cag acc ttc acc gcc agc Lys Lys Ser Thr Arg Ser Tyr Val Ala Ser Gln Thr Phe Thr Ala Ser 610 615 620	1872
ttc cgc cca ctg cat ggg ctg gac cgg tgg atg agt tac ctc gtg tgg Phe Ala Pro Leu His Gly Leu Asp Arg Trp Met Ser Tyr Leu Val Trp 625 630 635 640	1920
gtg acc gtt ttc gcc gcc aaa tat gcg gag tcg tac ttc ttc ctg atc Val Thr Val Phe Ala Ala Lys Tyr Ala Glu Ser Tyr Phe Phe Leu Ile 645 650 655	1968
ctt tcg ctc cgc gat ccg att cgc atc ctg tcg acg acc tcg atg cgc Leu Ser Leu Arg Asp Pro Ile Arg Ile Leu Ser Thr Thr Ser Met Arg 660 665 670	2016
tgc acg ggc gag tac tgg tgg gga aac aag atc tgc aag gtc caa ccg Cys Thr Gly Glu Tyr Trp Trp Gly Asn Lys Ile Cys Lys Val Gln Pro 675 680 685	2064
aag atc gtc ttg ggc ctc atg atc gcc acg gac ttc atc ctg ttc ttc Lys Ile Val Leu Gly Leu Met Ile Ala Thr Asp Phe Ile Leu Phe Phe 690 695 700	2112
ctg gac acc tac ctc tgg tac atc gtc gtg aac acg gtt ttc agc gtc Leu Asp Thr Tyr Leu Trp Tyr Ile Val Val Asn Thr Val Phe Ser Val 705 710 715 720	2160
gga aaa tcg ttc tac ctg ggc atc tcc att ctc acc ccg tgg cgc aac Gly Lys Ser Phe Tyr Leu Gly Ile Ser Ile Leu Thr Pro Trp Arg Asn 725 730 735	2208
atc ttt acc cgg ctg ccc aag cgc ata tac agc aag atc ctg gcc acc Ile Phe Thr Arg Leu Pro Lys Arg Ile Tyr Ser Lys Ile Leu Ala Thr 740 745 750	2256
acc gac atg gag atc aag tac aaa ccc aag gtc ctc atc agc cag atc Thr Asp Met Glu Ile Lys Tyr Lys Pro Lys Val Leu Ile Ser Gln Ile 755 760 765	2304
tgg aat gcg ata atc ata tcc atg tat cgc gag cat ctt ctg gcg atc Trp Asn Ala Ile Ile Ile Ser Met Tyr Arg Glu His Leu Leu Ala Ile 770 775 780	2352
gac cat gtc caa aag ctg ctg tat cat cag gtg ccg agc gag atc gag Asp His Val Gln Lys Leu Leu Tyr His Gln Val Pro Ser Glu Ile Glu 785 790 795 800	2400
ggg aag cgc acc ttg cgc gct ccg acc ttc ttc gtt tcg caa gac gat Gly Lys Arg Thr Leu Arg Ala Pro Thr Phe Phe Val Ser Gln Asp Asp 805 810 815	2448
aac aac ttc gag act gag ttc ttc cca agg gat tcc gag gcc gag cgc Asn Asn Phe Glu Thr Glu Phe Phe Pro Arg Asp Ser Glu Ala Glu Arg 820 825 830	2496
cgg atc agc ttt ttc gcg cag agc ctt tcc acc ccc atc ccc gag ccg Arg Ile Ser Phe Phe Ala Gln Ser Leu Ser Thr Pro Ile Pro Glu Pro 835 840 845	2544
ctg ccg gtg gac aac atg ccc acg ttc acg gtt ctg acc ccg cat tac Leu Pro Val Asp Asn Met Pro Thr Phe Thr Val Leu Thr Pro His Tyr 850 855 860	2592
gcc gag cgc atc ttg ctg agc ctg cgg gaa atc atc cgc gag gac gac Ala Glu Arg Ile Leu Leu Ser Leu Arg Glu Ile Ile Arg Glu Asp Asp 865 870 875 880	2640
cag ttc agc cgc gtg acc ctc ctg gag tat ctg aag caa ctc cat ccc Gln Phe Ser Arg Val Thr Leu Leu Glu Tyr Leu Lys Gln Leu His Pro 885 890 895	2688
gtg gag tgg gat tgc ttc gtc aaa gac acc aag atc ctg gcc gag gaa Val Glu Trp Asp Cys Phe Val Lys Asp Thr Lys Ile Leu Ala Glu Glu 900 905 910	2736

-continued

acc gcc gct tac gaa aac aac gag gac gag cct gag aag gag gac gcg Thr Ala Ala Tyr Glu Asn Asn Glu Asp Glu Pro Glu Lys Glu Asp Ala 915 920 925	2784
ctc aaa agt cag atc gac gac ttg ccg ttc tac tgc att gga ttc aaa Leu Lys Ser Gln Ile Asp Asp Leu Pro Phe Tyr Cys Ile Gly Phe Lys 930 935 940	2832
tcg gcg gca ccg gag tac acg ctt ccg acc ccg atc tgg gcg agc ctc Ser Ala Ala Pro Glu Tyr Thr Leu Arg Thr Arg Ile Trp Ala Ser Leu 945 950 955 960	2880
cgc tcg cag acc ctc tat ccg acc atc tcg gga ttc atg aat tac tcc Arg Ser Gln Thr Leu Tyr Arg Thr Ile Ser Gly Phe Met Asn Tyr Ser 965 970 975	2928
ccg gca atc aag ctg ctg tat ccg gtg gag aac ccg gag atc gtg caa Arg Ala Ile Lys Leu Leu Tyr Arg Val Glu Asn Pro Glu Ile Val Gln 980 985 990	2976
atg ttc ggc ggc aat gcc gac ggc ctg gag cgc gag ctt gag aag atg Met Phe Gly Gly Asn Ala Asp Gly Leu Glu Arg Glu Leu Glu Lys Met 995 1000 1005	3024
gcg agg cgc aag ttc aag ttc ctc gtg agt atg cag cgc ctg gca Ala Arg Arg Lys Phe Lys Phe Leu Val Ser Met Gln Arg Leu Ala 1010 1015 1020	3069
aag ttc aag ttc ctg gag aac gcc gag ttc ctg ctt ccg gcg tat Lys Phe Lys Phe Leu Glu Asn Ala Glu Phe Leu Leu Arg Ala Tyr 1025 1030 1035	3114
ccg gac ctc cag atc gcc tac ctg gac gaa gaa ccg ccc ctg aat Pro Asp Leu Gln Ile Ala Tyr Leu Asp Glu Glu Pro Pro Leu Asn 1040 1045 1050	3159
gag ggc gag gag cca cgc atc tat tcc gcg ctc atc gac ggc cac Glu Gly Glu Glu Pro Arg Ile Tyr Ser Ala Leu Ile Asp Gly His 1055 1060 1065	3204
tgc gag att aat ggc cgc ccg cgc ccg aag ttc agg gtc cag ctc Cys Glu Ile Asn Gly Arg Arg Arg Pro Lys Phe Arg Val Gln Leu 1070 1075 1080	3249
tcc ggc aac ccg att ttg ggc gat ggg aaa tcg gac aac cag aat Ser Gly Asn Pro Ile Leu Gly Asp Gly Lys Ser Asp Asn Gln Asn 1085 1090 1095	3294
cat gcg ctc atc ttc tat ccg ggc gag tac att cag ctg atc gac His Ala Leu Ile Phe Tyr Arg Gly Glu Tyr Ile Gln Leu Ile Asp 1100 1105 1110	3339
gcc aac caa gat aac tat ctt gag gag tgc ctg aag atc cgc tcg Ala Asn Gln Asp Asn Tyr Leu Glu Glu Cys Leu Lys Ile Arg Ser 1115 1120 1125	3384
gtg ctg gcg gag ttt gag gaa ctc ggt att gaa cag ata cac ccg Val Leu Ala Glu Phe Glu Glu Leu Gly Ile Glu Gln Ile His Pro 1130 1135 1140	3429
tat acc ccc gga ctc aag tac gag gac cag tcg acg aat cat cca Tyr Thr Pro Gly Leu Lys Tyr Glu Asp Gln Ser Thr Asn His Pro 1145 1150 1155	3474
gtg gct atc gtc ggt gcc cgc gag tat atc ttc agc gag aac tcc Val Ala Ile Val Gly Ala Arg Glu Tyr Ile Phe Ser Glu Asn Ser 1160 1165 1170	3519
ggt gtc ctc ggc gac gtg gcg gct ggc aag gaa cag acc ttc ggc Gly Val Leu Gly Asp Val Ala Ala Gly Lys Glu Gln Thr Phe Gly 1175 1180 1185	3564
acc ctc ttc gcc ccg acc ctc gcg cag atc ggc ggc aag ctg cac Thr Leu Phe Ala Arg Thr Leu Ala Gln Ile Gly Gly Lys Leu His 1190 1195 1200	3609

-continued

tac ggc cat ccg gac ttc atc aac gcg acc ttc atg acg acc cgg	3654
Tyr Gly His Pro Asp Phe Ile Asn Ala Thr Phe Met Thr Thr Arg	
1205 1210 1215	
ggt ggg gtc agc aag gcc cag aaa ggg ctg cat ctc aat gaa gat	3699
Gly Gly Val Ser Lys Ala Gln Lys Gly Leu His Leu Asn Glu Asp	
1220 1225 1230	
atc tac gcg ggg atg aac gct gtc ctc cgc ggt ggc cgc att aag	3744
Ile Tyr Ala Gly Met Asn Ala Val Leu Arg Gly Gly Arg Ile Lys	
1235 1240 1245	
cac tgc gag tat tac cag tgc ggc aag ggt cgc gac ttg ggc ttt	3789
His Cys Glu Tyr Tyr Gln Cys Gly Lys Gly Arg Asp Leu Gly Phe	
1250 1255 1260	
ggc acc atc ctc aac ttc acg acg aaa atc ggc gct ggc atg ggc	3834
Gly Thr Ile Leu Asn Phe Thr Thr Lys Ile Gly Ala Gly Met Gly	
1265 1270 1275	
gag caa atg ctg agt cgg gag tac tat tac ctg ggc acc cag ctg	3879
Glu Gln Met Leu Ser Arg Glu Tyr Tyr Tyr Leu Gly Thr Gln Leu	
1280 1285 1290	
cca atc gac cgc ttc ctc acc ttc tac tat gcg cat ccc ggc ttc	3924
Pro Ile Asp Arg Phe Leu Thr Phe Tyr Tyr Ala His Pro Gly Phe	
1295 1300 1305	
cat ctg aat aac ctg ttc ata cag ctg tcg ctg cag atg ttc atg	3969
His Leu Asn Asn Leu Phe Ile Gln Leu Ser Leu Gln Met Phe Met	
1310 1315 1320	
ctg act ctg gtc aac ctc cac gca ctc gcg cat gag agc att ttg	4014
Leu Thr Leu Val Asn Leu His Ala Leu Ala His Glu Ser Ile Leu	
1325 1330 1335	
tgc gtg tat gac agg gat aaa ccg atc acc gac gtc ctc tat cct	4059
Cys Val Tyr Asp Arg Asp Lys Pro Ile Thr Asp Val Leu Tyr Pro	
1340 1345 1350	
atc ggt tgc tat aac ttc cat ccg gcg atc gac tgg gtt cgc cgc	4104
Ile Gly Cys Tyr Asn Phe His Pro Ala Ile Asp Trp Val Arg Arg	
1355 1360 1365	
tac acc ctg agc atc ttt atc gtg ttc tgg ata gcg ttc gtc ccg	4149
Tyr Thr Leu Ser Ile Phe Ile Val Phe Trp Ile Ala Phe Val Pro	
1370 1375 1380	
atc gtc gtc cag gag ctg atc gag agg ggc ctg tgg aag gcc acg	4194
Ile Val Val Gln Glu Leu Ile Glu Arg Gly Leu Trp Lys Ala Thr	
1385 1390 1395	
cag cgg ttc ttc agg cac atc ttg tcg ctc agc ccg atg ttc gaa	4239
Gln Arg Phe Phe Arg His Ile Leu Ser Leu Ser Pro Met Phe Glu	
1400 1405 1410	
gtg ttc gcg ggt cag atc tat tcc agc gcg ctc ctg agc gac att	4284
Val Phe Ala Gly Gln Ile Tyr Ser Ser Ala Leu Leu Ser Asp Ile	
1415 1420 1425	
gcc gtg ggc ggc gcc cgg tac atc tcc acg ggc cgc ggt ttc gcg	4329
Ala Val Gly Gly Ala Arg Tyr Ile Ser Thr Gly Arg Gly Phe Ala	
1430 1435 1440	
acc tcc cgc atc ccg ttc tcg atc ctc tat tcg cgc ttc gcg ggc	4374
Thr Ser Arg Ile Pro Phe Ser Ile Leu Tyr Ser Arg Phe Ala Gly	
1445 1450 1455	
agc gcg atc tac atg ggt tcc cgg agc atg ctc atg ctc ctg ttc	4419
Ser Ala Ile Tyr Met Gly Ser Arg Ser Met Leu Met Leu Leu Phe	
1460 1465 1470	
ggc acc gtc gcc cac tgg cag gct ccc ctg ctt tgg ttc tgg gcg	4464
Gly Thr Val Ala His Trp Gln Ala Pro Leu Leu Trp Phe Trp Ala	
1475 1480 1485	

-continued

agc ctt agc gcg ctg atc ttc gcc cgg ttc atc ttc aat ccg cat	4509
Ser Leu Ser Ala Leu Ile Phe Ala Pro Phe Ile Phe Asn Pro His	
1490 1495 1500	
cag ttc gcc tgg gag gac ttc ttc ctc gac tat agg gac tac atc	4554
Gln Phe Ala Trp Glu Asp Phe Phe Leu Asp Tyr Arg Asp Tyr Ile	
1505 1510 1515	
cgc tgg ctg tcc cgc gga aac aac aag tat cac cgc aac tcg tgg	4599
Arg Trp Leu Ser Arg Gly Asn Asn Lys Tyr His Arg Asn Ser Trp	
1520 1525 1530	
atc ggt tac gtg cgc atg tgc cgg tcc cgg gtg acg ggt ttc aag	4644
Ile Gly Tyr Val Arg Met Ser Arg Ser Arg Val Thr Gly Phe Lys	
1535 1540 1545	
cgc aag ctc gtc ggc gat gag tgc gag aag agt gcc ggg gac gcc	4689
Arg Lys Leu Val Gly Asp Glu Ser Glu Lys Ser Ala Gly Asp Ala	
1550 1555 1560	
tcc cgc gcc cat cgc acc aac ctc atc atg gcc gag atc atc ccg	4734
Ser Arg Ala His Arg Thr Asn Leu Ile Met Ala Glu Ile Ile Pro	
1565 1570 1575	
tgc gcc atc tac gcg gca ggc tgc ttc atc gcc ttc acc ttt atc	4779
Cys Ala Ile Tyr Ala Ala Gly Cys Phe Ile Ala Phe Thr Phe Ile	
1580 1585 1590	
aac gcg cag acc ggc gtc aag acc acc gat gag gac cgc gtg aac	4824
Asn Ala Gln Thr Gly Val Lys Thr Thr Asp Glu Asp Arg Val Asn	
1595 1600 1605	
tcc acg ctg cgc atc atc atc tgt acc ctg gca ccg atc gtc ata	4869
Ser Thr Leu Arg Ile Ile Ile Cys Thr Leu Ala Pro Ile Val Ile	
1610 1615 1620	
gac atc ggg gtg ctg ttc ttc tgc atg ggc ttg agc tgt tgc tgc	4914
Asp Ile Gly Val Leu Phe Phe Cys Met Gly Leu Ser Cys Cys Ser	
1625 1630 1635	
gga ccg ctg ctc ggc atg tgc tgc aag aaa act gcc agc gtc atg	4959
Gly Pro Leu Leu Gly Met Cys Cys Lys Lys Thr Gly Ser Val Met	
1640 1645 1650	
gcc ggc att gcc cat ggc atc gcc gtc gtc gtc cat att gtc ttt	5004
Ala Gly Ile Ala His Gly Ile Ala Val Val Val His Ile Val Phe	
1655 1660 1665	
ttc atc gtc atg tgg gtg ctc gaa ggc ttc tcc ttc gtc cgc atg	5049
Phe Ile Val Met Trp Val Leu Glu Gly Phe Ser Phe Val Arg Met	
1670 1675 1680	
ctg atc ggt gtc gtg acg tgc atc cag tgt cag cgg ctt atc ttt	5094
Leu Ile Gly Val Val Thr Cys Ile Gln Cys Gln Arg Leu Ile Phe	
1685 1690 1695	
cat tgc atg acc gtg ctc ctc ctc acg cgg gaa ttc aag aat gac	5139
His Cys Met Thr Val Leu Leu Leu Thr Arg Glu Phe Lys Asn Asp	
1700 1705 1710	
cac gcc aac acc gcg ttc tgg acc gcc aag tgg tat tcg acg gga	5184
His Ala Asn Thr Ala Phe Trp Thr Gly Lys Trp Tyr Ser Thr Gly	
1715 1720 1725	
ctc ggt tat atg gcc tgg acg cag cca acg cgc gag ctg acg gca	5229
Leu Gly Tyr Met Ala Trp Thr Gln Pro Thr Arg Glu Leu Thr Ala	
1730 1735 1740	
aag gtt atc gag ctg agc gaa ttc gcg gct gac ttc gtg ctg ggc	5274
Lys Val Ile Glu Leu Ser Glu Phe Ala Ala Asp Phe Val Leu Gly	
1745 1750 1755	
cat gtc atc ctc atc ttc cag ctc ccg gtt atc tgc atc ccg aag	5319
His Val Ile Leu Ile Phe Gln Leu Pro Val Ile Cys Ile Pro Lys	
1760 1765 1770	

-continued

```

atc gac aaa ttc cac tcg att atg ctc ttt tgg ctg aag ccg tcg 5364
Ile Asp Lys Phe His Ser Ile Met Leu Phe Trp Leu Lys Pro Ser
1775 1780 1785

cgc caa atc cgc cct ccg atc tac tcc ctg aaa cag gcg cgc ctg 5409
Arg Gln Ile Arg Pro Pro Ile Tyr Ser Leu Lys Gln Ala Arg Leu
1790 1795 1800

cgc aag cgc atg gtc cgg cgc tac tgc tcc ctc tac ttc ctg gtg 5454
Arg Lys Arg Met Val Arg Arg Tyr Cys Ser Leu Tyr Phe Leu Val
1805 1810 1815

ctg atc atc ttc gcg ggt tgc atc gtg ggt ccc gcc gtg gcg tcg 5499
Leu Ile Ile Phe Ala Gly Cys Ile Val Gly Pro Ala Val Ala Ser
1820 1825 1830

gct cac gtg ccg aaa gac ctg ggc tcc ggg ctg acc ggc acc ttt 5544
Ala His Val Pro Lys Asp Leu Gly Ser Gly Leu Thr Gly Thr Phe
1835 1840 1845

cat aac ctg gtg cag ccg cgc aac gtc agc aat aat gac acg gcc 5589
His Asn Leu Val Gln Pro Arg Asn Val Ser Asn Asn Asp Thr Gly
1850 1855 1860

tcg caa atg tcc acc tat aag tcg cat tac tac acc cat acc ccg 5634
Ser Gln Met Ser Thr Tyr Lys Ser His Tyr Tyr Thr His Thr Pro
1865 1870 1875

tcg ctc aaa acc tgg tcg acc att aaa 5661
Ser Leu Lys Thr Trp Ser Thr Ile Lys
1880 1885

```

```

<210> SEQ ID NO 8
<211> LENGTH: 1887
<212> TYPE: PRT
<213> ORGANISM: Artificial sequence
<220> FEATURE:
<223> OTHER INFORMATION: Synthetic Construct

```

```

<400> SEQUENCE: 8

```

```

Met Ser Tyr Asn Asp Pro Asn Leu Asn Gly Gln Tyr Tyr Ser Asn Gly
1 5 10 15
Asp Gly Thr Gly Asp Gly Asn Tyr Pro Thr Tyr Gln Val Thr Gln Asp
20 25 30
Gln Ser Ala Tyr Asp Glu Tyr Gly Gln Pro Ile Tyr Thr Gln Asn Gln
35 40 45
Leu Asp Asp Gly Tyr Tyr Asp Pro Asn Glu Gln Tyr Val Asp Gly Thr
50 55 60
Gln Phe Pro Gln Gly Gln Asp Pro Ser Gln Asp Gln Gly Pro Tyr Asn
65 70 75 80
Asn Asp Ala Ser Tyr Tyr Asn Gln Pro Pro Asn Met Met Asn Pro Ser
85 90 95
Ser Gln Asp Gly Glu Asn Phe Ser Asp Phe Ser Ser Tyr Gly Pro Pro
100 105 110
Ser Gly Thr Tyr Pro Asn Asp Gln Tyr Thr Pro Ser Gln Met Ser Tyr
115 120 125
Pro Asp Gln Asp Gly Ser Ser Gly Ala Ser Thr Pro Tyr Gly Asn Gly
130 135 140
Val Val Asn Gly Asn Gly Gln Tyr Tyr Asp Pro Asn Ala Ile Glu Met
145 150 155 160
Ala Leu Pro Asn Asp Pro Tyr Pro Ala Trp Thr Ala Asp Pro Gln Ser
165 170 175

```


-continued

Pro	Leu	Pro	Ile	Glu	Gln	Ile	Glu	Asp	Ile	Phe	Ile	Asp	Leu	Thr	Asn
			180					185					190		
Lys	Phe	Gly	Phe	Gln	Arg	Asp	Ser	Met	Arg	Asn	Met	Phe	Asp	His	Phe
		195					200					205			
Met	Thr	Leu	Leu	Asp	Ser	Arg	Ser	Ser	Arg	Met	Ser	Pro	Glu	Gln	Ala
	210					215					220				
Leu	Leu	Ser	Leu	His	Ala	Asp	Tyr	Ile	Gly	Gly	Asp	Thr	Ala	Asn	Tyr
225					230					235					240
Lys	Lys	Trp	Tyr	Phe	Ala	Ala	Gln	Leu	Asp	Met	Asp	Asp	Glu	Ile	Gly
				245					250					255	
Phe	Arg	Asn	Met	Lys	Leu	Gly	Lys	Leu	Ser	Arg	Lys	Ala	Arg	Lys	Ala
			260					265					270		
Lys	Lys	Lys	Asn	Lys	Lys	Ala	Met	Gln	Glu	Asp	Glu	Asp	Thr	Glu	Glu
		275					280					285			
Thr	Leu	Asn	Gln	Ile	Glu	Gly	Asp	Asn	Ser	Leu	Glu	Ala	Ala	Asp	Phe
	290					295					300				
Arg	Trp	Lys	Ser	Lys	Met	Asn	Gln	Leu	Ser	Pro	Phe	Glu	Met	Val	Arg
305					310					315					320
Gln	Ile	Ala	Leu	Phe	Leu	Cys	Trp	Gly	Glu	Ala	Asn	Gln	Val	Arg	
				325				330						335	
Phe	Thr	Pro	Glu	Cys	Leu	Cys	Phe	Ile	Tyr	Lys	Cys	Ala	Ser	Asp	Tyr
			340					345					350		
Leu	Asp	Ser	Ala	Gln	Cys	Gln	Gln	Arg	Pro	Asp	Pro	Leu	Pro	Glu	Gly
		355					360					365			
Asp	Phe	Leu	Asn	Arg	Val	Ile	Thr	Pro	Leu	Tyr	Arg	Phe	Ile	Arg	Ser
	370					375					380				
Gln	Val	Tyr	Glu	Ile	Val	Asp	Gly	Arg	Tyr	Val	Lys	Ser	Glu	Lys	Asp
385					390					395					400
His	Asn	Lys	Val	Ile	Gly	Tyr	Asp	Asp	Val	Asn	Gln	Leu	Phe	Trp	Tyr
				405					410					415	
Pro	Glu	Gly	Ile	Ala	Lys	Ile	Val	Met	Glu	Asp	Gly	Thr	Arg	Leu	Ile
			420					425					430		
Asp	Leu	Pro	Ala	Glu	Glu	Arg	Tyr	Leu	Lys	Leu	Gly	Glu	Ile	Pro	Trp
		435					440					445			
Asp	Asp	Val	Phe	Phe	Lys	Thr	Tyr	Lys	Glu	Thr	Arg	Ser	Trp	Leu	His
	450					455					460				
Leu	Val	Thr	Asn	Phe	Asn	Arg	Ile	Trp	Ile	Met	His	Val	Tyr	Trp	Met
465					470					475					480
Tyr	Cys	Ala	Tyr	Asn	Ala	Pro	Thr	Phe	Tyr	Thr	His	Asn	Tyr	Gln	Gln
				485					490					495	
Leu	Val	Asp	Asn	Gln	Pro	Leu	Ala	Ala	Tyr	Lys	Trp	Ala	Thr	Ala	Ala
			500					505					510		
Leu	Gly	Gly	Thr	Val	Ala	Ser	Leu	Ile	Gln	Val	Ala	Ala	Thr	Leu	Cys
		515					520					525			
Glu	Trp	Ser	Phe	Val	Pro	Arg	Lys	Trp	Ala	Gly	Ala	Gln	His	Leu	Ser
	530					535					540				
Arg	Arg	Phe	Trp	Phe	Leu	Cys	Val	Ile	Met	Gly	Ile	Asn	Leu	Gly	Pro
545					550					555					560
Val	Ile	Phe	Val	Phe	Ala	Tyr	Asp	Lys	Asp	Thr	Val	Tyr	Ser	Thr	Ala
				565					570					575	
Ala	His	Val	Val	Gly	Ala	Val	Met	Phe	Phe	Val	Ala	Val	Ala	Thr	Leu

-continued

580					585					590					
Val	Phe	Phe	Ser	Val	Met	Pro	Leu	Gly	Gly	Leu	Phe	Thr	Ser	Tyr	Met
	595						600					605			
Lys	Lys	Ser	Thr	Arg	Ser	Tyr	Val	Ala	Ser	Gln	Thr	Phe	Thr	Ala	Ser
	610					615					620				
Phe	Ala	Pro	Leu	His	Gly	Leu	Asp	Arg	Trp	Met	Ser	Tyr	Leu	Val	Trp
625					630					635					640
Val	Thr	Val	Phe	Ala	Ala	Lys	Tyr	Ala	Glu	Ser	Tyr	Phe	Phe	Leu	Ile
				645					650					655	
Leu	Ser	Leu	Arg	Asp	Pro	Ile	Arg	Ile	Leu	Ser	Thr	Thr	Ser	Met	Arg
			660					665					670		
Cys	Thr	Gly	Glu	Tyr	Trp	Trp	Gly	Asn	Lys	Ile	Cys	Lys	Val	Gln	Pro
	675						680					685			
Lys	Ile	Val	Leu	Gly	Leu	Met	Ile	Ala	Thr	Asp	Phe	Ile	Leu	Phe	Phe
	690					695					700				
Leu	Asp	Thr	Tyr	Leu	Trp	Tyr	Ile	Val	Val	Asn	Thr	Val	Phe	Ser	Val
705					710					715					720
Gly	Lys	Ser	Phe	Tyr	Leu	Gly	Ile	Ser	Ile	Leu	Thr	Pro	Trp	Arg	Asn
				725					730					735	
Ile	Phe	Thr	Arg	Leu	Pro	Lys	Arg	Ile	Tyr	Ser	Lys	Ile	Leu	Ala	Thr
			740					745					750		
Thr	Asp	Met	Glu	Ile	Lys	Tyr	Lys	Pro	Lys	Val	Leu	Ile	Ser	Gln	Ile
		755					760					765			
Trp	Asn	Ala	Ile	Ile	Ile	Ser	Met	Tyr	Arg	Glu	His	Leu	Leu	Ala	Ile
	770					775					780				
Asp	His	Val	Gln	Lys	Leu	Leu	Tyr	His	Gln	Val	Pro	Ser	Glu	Ile	Glu
785					790					795					800
Gly	Lys	Arg	Thr	Leu	Arg	Ala	Pro	Thr	Phe	Phe	Val	Ser	Gln	Asp	Asp
				805					810					815	
Asn	Asn	Phe	Glu	Thr	Glu	Phe	Phe	Pro	Arg	Asp	Ser	Glu	Ala	Glu	Arg
			820					825					830		
Arg	Ile	Ser	Phe	Phe	Ala	Gln	Ser	Leu	Ser	Thr	Pro	Ile	Pro	Glu	Pro
		835					840					845			
Leu	Pro	Val	Asp	Asn	Met	Pro	Thr	Phe	Thr	Val	Leu	Thr	Pro	His	Tyr
	850					855					860				
Ala	Glu	Arg	Ile	Leu	Leu	Ser	Leu	Arg	Glu	Ile	Ile	Arg	Glu	Asp	Asp
865					870					875					880
Gln	Phe	Ser	Arg	Val	Thr	Leu	Leu	Glu	Tyr	Leu	Lys	Gln	Leu	His	Pro
				885					890					895	
Val	Glu	Trp	Asp	Cys	Phe	Val	Lys	Asp	Thr	Lys	Ile	Leu	Ala	Glu	Glu
		900						905					910		
Thr	Ala	Ala	Tyr	Glu	Asn	Asn	Glu	Asp	Glu	Pro	Glu	Lys	Glu	Asp	Ala
		915					920					925			
Leu	Lys	Ser	Gln	Ile	Asp	Asp	Leu	Pro	Phe	Tyr	Cys	Ile	Gly	Phe	Lys
	930					935					940				
Ser	Ala	Ala	Pro	Glu	Tyr	Thr	Leu	Arg	Thr	Arg	Ile	Trp	Ala	Ser	Leu
945					950					955					960
Arg	Ser	Gln	Thr	Leu	Tyr	Arg	Thr	Ile	Ser	Gly	Phe	Met	Asn	Tyr	Ser
				965					970					975	
Arg	Ala	Ile	Lys	Leu	Leu	Tyr	Arg	Val	Glu	Asn	Pro	Glu	Ile	Val	Gln
			980					985					990		

-continued

Met	Phe	Gly	Gly	Asn	Ala	Asp	Gly	Leu	Glu	Arg	Glu	Leu	Glu	Lys	Met
	995						1000					1005			
Ala	Arg	Arg	Lys	Phe	Lys	Phe	Leu	Val	Ser	Met	Gln	Arg	Leu	Ala	
1010						1015					1020				
Lys	Phe	Lys	Phe	Leu	Glu	Asn	Ala	Glu	Phe	Leu	Leu	Arg	Ala	Tyr	
1025						1030					1035				
Pro	Asp	Leu	Gln	Ile	Ala	Tyr	Leu	Asp	Glu	Glu	Pro	Pro	Leu	Asn	
1040						1045					1050				
Glu	Gly	Glu	Glu	Pro	Arg	Ile	Tyr	Ser	Ala	Leu	Ile	Asp	Gly	His	
1055						1060					1065				
Cys	Glu	Ile	Asn	Gly	Arg	Arg	Arg	Pro	Lys	Phe	Arg	Val	Gln	Leu	
1070						1075					1080				
Ser	Gly	Asn	Pro	Ile	Leu	Gly	Asp	Gly	Lys	Ser	Asp	Asn	Gln	Asn	
1085						1090					1095				
His	Ala	Leu	Ile	Phe	Tyr	Arg	Gly	Glu	Tyr	Ile	Gln	Leu	Ile	Asp	
1100						1105					1110				
Ala	Asn	Gln	Asp	Asn	Tyr	Leu	Glu	Glu	Cys	Leu	Lys	Ile	Arg	Ser	
1115						1120					1125				
Val	Leu	Ala	Glu	Phe	Glu	Glu	Leu	Gly	Ile	Glu	Gln	Ile	His	Pro	
1130						1135					1140				
Tyr	Thr	Pro	Gly	Leu	Lys	Tyr	Glu	Asp	Gln	Ser	Thr	Asn	His	Pro	
1145						1150					1155				
Val	Ala	Ile	Val	Gly	Ala	Arg	Glu	Tyr	Ile	Phe	Ser	Glu	Asn	Ser	
1160						1165					1170				
Gly	Val	Leu	Gly	Asp	Val	Ala	Ala	Gly	Lys	Glu	Gln	Thr	Phe	Gly	
1175						1180					1185				
Thr	Leu	Phe	Ala	Arg	Thr	Leu	Ala	Gln	Ile	Gly	Gly	Lys	Leu	His	
1190						1195					1200				
Tyr	Gly	His	Pro	Asp	Phe	Ile	Asn	Ala	Thr	Phe	Met	Thr	Thr	Arg	
1205						1210					1215				
Gly	Gly	Val	Ser	Lys	Ala	Gln	Lys	Gly	Leu	His	Leu	Asn	Glu	Asp	
1220						1225					1230				
Ile	Tyr	Ala	Gly	Met	Asn	Ala	Val	Leu	Arg	Gly	Gly	Arg	Ile	Lys	
1235						1240					1245				
His	Cys	Glu	Tyr	Tyr	Gln	Cys	Gly	Lys	Gly	Arg	Asp	Leu	Gly	Phe	
1250						1255					1260				
Gly	Thr	Ile	Leu	Asn	Phe	Thr	Thr	Lys	Ile	Gly	Ala	Gly	Met	Gly	
1265						1270					1275				
Glu	Gln	Met	Leu	Ser	Arg	Glu	Tyr	Tyr	Tyr	Leu	Gly	Thr	Gln	Leu	
1280						1285					1290				
Pro	Ile	Asp	Arg	Phe	Leu	Thr	Phe	Tyr	Tyr	Ala	His	Pro	Gly	Phe	
1295						1300					1305				
His	Leu	Asn	Asn	Leu	Phe	Ile	Gln	Leu	Ser	Leu	Gln	Met	Phe	Met	
1310						1315					1320				
Leu	Thr	Leu	Val	Asn	Leu	His	Ala	Leu	Ala	His	Glu	Ser	Ile	Leu	
1325						1330					1335				
Cys	Val	Tyr	Asp	Arg	Asp	Lys	Pro	Ile	Thr	Asp	Val	Leu	Tyr	Pro	
1340						1345					1350				
Ile	Gly	Cys	Tyr	Asn	Phe	His	Pro	Ala	Ile	Asp	Trp	Val	Arg	Arg	
1355						1360					1365				

-continued

1745	1750	1755	
His Val Ile Leu Ile Phe Gln	Leu Pro Val Ile Cys	Ile Pro Lys	
1760	1765	1770	
Ile Asp Lys Phe His Ser Ile	Met Leu Phe Trp Leu	Lys Pro Ser	
1775	1780	1785	
Arg Gln Ile Arg Pro Pro Ile	Tyr Ser Leu Lys Gln	Ala Arg Leu	
1790	1795	1800	
Arg Lys Arg Met Val Arg Arg	Tyr Cys Ser Leu Tyr	Phe Leu Val	
1805	1810	1815	
Leu Ile Ile Phe Ala Gly Cys	Ile Val Gly Pro Ala	Val Ala Ser	
1820	1825	1830	
Ala His Val Pro Lys Asp Leu	Gly Ser Gly Leu Thr	Gly Thr Phe	
1835	1840	1845	
His Asn Leu Val Gln Pro Arg	Asn Val Ser Asn Asn	Asp Thr Gly	
1850	1855	1860	
Ser Gln Met Ser Thr Tyr Lys	Ser His Tyr Tyr Thr	His Thr Pro	
1865	1870	1875	
Ser Leu Lys Thr Trp Ser Thr	Ile Lys		
1880	1885		
<210> SEQ ID NO 9			
<211> LENGTH: 5679			
<212> TYPE: DNA			
<213> ORGANISM: Artificial sequence			
<220> FEATURE:			
<223> OTHER INFORMATION: Synthetic construct			
<220> FEATURE:			
<221> NAME/KEY: CDS			
<222> LOCATION: (1)..(5679)			
<223> OTHER INFORMATION: Candida albicans FKS1 (codon optimized)			
<400> SEQUENCE: 9			
atg agc tac aac gac aac aat aat cac tat tac gac ccg aat cag caa			48
Met Ser Tyr Asn Asp Asn Asn Asn His Tyr Tyr Asp Pro Asn Gln Gln			
1 5 10 15			
ggc gga atg ccc cct cat caa ggc gga gag ggg tat tac cag cag cag			96
Gly Gly Met Pro His Gln Gly Gly Glu Gly Tyr Tyr Gln Gln Gln			
20 25 30			
tat gat gac atg ggc caa caa ccg cac caa cag gac tac tac gat ccg			144
Tyr Asp Asp Met Gly Gln Gln Pro His Gln Gln Asp Tyr Tyr Asp Pro			
35 40 45			
aat gct cag tat cag cag cag ccg tat gac atg gac ggt tat cag gac			192
Asn Ala Gln Tyr Gln Gln Gln Pro Tyr Asp Met Asp Gly Tyr Gln Asp			
50 55 60			
caa cgc aac tac ggc ggt cag ccg atg aat gcg caa ggc tac aac gcc			240
Gln Ala Asn Tyr Gly Gly Gln Pro Met Asn Ala Gln Gly Tyr Asn Ala			
65 70 75 80			
gat ccc gag gcc ttt tcc gac ttc agc tac ggt ggc cag acc ccg ggt			288
Asp Pro Glu Ala Phe Ser Asp Phe Ser Tyr Gly Gly Gln Thr Pro Gly			
85 90 95			
acc cca ggc tac gac cag tac ggt acc cag tac acc ccg agc cag atg			336
Thr Pro Gly Tyr Asp Gln Tyr Gly Thr Gln Tyr Thr Pro Ser Gln Met			
100 105 110			
tcc tat ggc ggc gac ccc cgg tcc tcg ggt gca agc act ccc att tat			384
Ser Tyr Gly Gly Asp Pro Arg Ser Ser Gly Ala Ser Thr Pro Ile Tyr			
115 120 125			
ggc ggc cag ggc cag ggt tac gac cct acc cag ttc aac atg tcc tcc			432

-continued

Gly	Gly	Gln	Gly	Gln	Gly	Tyr	Asp	Pro	Thr	Gln	Phe	Asn	Met	Ser	Ser	
aac	ctc	ccc	tac	ccc	gcc	tgg	agc	gcg	gac	ccc	cag	gcc	ccg	atc	aag	480
Asn	Leu	Pro	Tyr	Pro	Ala	Trp	Ser	Ala	Asp	Pro	Gln	Ala	Pro	Ile	Lys	
145				150						155					160	
ata	gag	cat	atc	gaa	gat	atc	ttc	atc	gac	ctc	acc	aac	aag	ttc	ggg	528
Ile	Glu	His	Ile	Glu	Asp	Ile	Phe	Ile	Asp	Leu	Thr	Asn	Lys	Phe	Gly	
				165					170					175		
ttc	cag	cgg	gac	tcg	atg	cgc	aat	atg	ttc	gac	tac	ttc	atg	acg	ctg	576
Phe	Gln	Arg	Asp	Ser	Met	Arg	Asn	Met	Phe	Asp	Tyr	Phe	Met	Thr	Leu	
				180				185						190		
ttg	gat	tcc	cgc	agc	tcg	cgc	atg	agc	ccg	gct	cag	gcc	ctc	ctc	agc	624
Leu	Asp	Ser	Arg	Ser	Ser	Arg	Met	Ser	Pro	Ala	Gln	Ala	Leu	Leu	Ser	
		195					200					205				
ctg	cat	gcg	gac	tat	att	ggc	ggg	gat	aac	gcg	aac	tac	cgg	aaa	tgg	672
Leu	His	Ala	Asp	Tyr	Ile	Gly	Gly	Asp	Asn	Ala	Asn	Tyr	Arg	Lys	Trp	
				210		215					220					
tac	ttc	agt	tcg	cag	caa	gat	ctc	gac	gac	tcg	ctg	ggc	ttc	cgc	aac	720
Tyr	Phe	Ser	Ser	Gln	Gln	Asp	Leu	Asp	Asp	Ser	Leu	Gly	Phe	Ala	Asn	
225					230					235					240	
atg	acc	ctg	ggg	aag	atc	gga	cgc	aag	gcc	cgg	aag	gcc	agc	aag	aaa	768
Met	Thr	Leu	Gly	Lys	Ile	Gly	Arg	Lys	Ala	Arg	Lys	Ala	Ser	Lys	Lys	
				245					250					255		
tcc	aag	aag	gcc	cgc	aaa	cgc	gcc	gag	gag	cac	ggc	cag	gac	gtg	gac	816
Ser	Lys	Lys	Ala	Arg	Lys	Ala	Ala	Glu	Glu	His	Gly	Gln	Asp	Val	Asp	
			260					265					270			
gcg	aac	gag	ctg	gaa	ggc	gac	tat	tcg	ctt	gag	gcc	gca	gag	atc	cgc	864
Ala	Asn	Glu	Leu	Glu	Gly	Asp	Tyr	Ser	Leu	Glu	Ala	Ala	Glu	Ile	Arg	
			275				280					285				
tgg	aag	gcc	aag	atg	aat	tcg	ctg	acc	ccg	gaa	gaa	cgc	gtg	cgg	gac	912
Trp	Lys	Ala	Lys	Met	Asn	Ser	Leu	Thr	Pro	Glu	Glu	Arg	Val	Arg	Asp	
		290				295						300				
ctg	gcg	ctg	tat	ctc	ctg	atc	tgg	ggc	gaa	gcg	aat	cag	gtc	agg	ttt	960
Leu	Ala	Leu	Tyr	Leu	Leu	Ile	Trp	Gly	Glu	Ala	Asn	Gln	Val	Arg	Phe	
				310						315					320	
acg	ccc	gag	tgt	ctc	tgc	tac	atc	tac	aaa	tcg	gcc	acc	gat	tat	ctg	1008
Thr	Pro	Glu	Cys	Leu	Cys	Tyr	Ile	Tyr	Lys	Ser	Ala	Thr	Asp	Tyr	Leu	
				325					330					335		
aat	tcg	ccc	ctt	tgc	cag	cag	cgc	cag	gaa	ccc	gtc	ccg	gaa	ggc	gac	1056
Asn	Ser	Pro	Leu	Cys	Gln	Gln	Arg	Gln	Glu	Pro	Val	Pro	Glu	Gly	Asp	
			340					345					350			
tat	ctg	aac	cgc	gtg	att	acc	ccc	ctt	tac	cgc	ttc	atc	cgg	tcc	caa	1104
Tyr	Leu	Asn	Arg	Val	Ile	Thr	Pro	Leu	Tyr	Arg	Phe	Ile	Arg	Ser	Gln	
				355			360						365			
gtc	tat	gag	atc	tac	gac	ggc	cgg	ttc	gtc	aag	cgc	gag	aag	gac	cac	1152
Val	Tyr	Glu	Ile	Tyr	Asp	Gly	Arg	Phe	Val	Lys	Arg	Glu	Lys	Asp	His	
				370		375					380					
aat	aag	gtc	att	gga	tac	gac	gac	gtg	aac	caa	ttg	ttc	tgg	tat	ccc	1200
Asn	Lys	Val	Ile	Gly	Tyr	Asp	Asp	Val	Asn	Gln	Leu	Phe	Trp	Tyr	Pro	
				385		390				395					400	
gag	ggc	atc	agc	cgc	atc	atc	ttc	gag	gat	ggc	acg	cgc	ctg	gtg	gac	1248
Glu	Gly	Ile	Ser	Arg	Ile	Ile	Phe	Glu	Asp	Gly	Thr	Arg	Leu	Val	Asp	
				405					410					415		
atc	ccg	cag	gag	gag	cgg	ttc	ttg	aag	ctc	gga	gag	gtc	gag	tgg	aaa	1296
Ile	Pro	Gln	Glu	Glu	Arg	Phe	Leu	Lys	Leu	Gly	Glu	Val	Glu	Trp	Lys	
				420				425						430		
aat	gtg	ttc	ttc	aag	acc	tat	aaa	gaa	atc	cgc	acc	tgg	ctg	cac	ttc	1344

-continued

Asn	Val	Phe	Phe	Lys	Thr	Tyr	Lys	Glu	Ile	Arg	Thr	Trp	Leu	His	Phe	
		435					440					445				
gtg	acc	aat	ttc	aac	cgg	atc	tgg	atc	atc	cat	ggc	acg	atc	tat	tgg	1392
Val	Thr	Asn	Phe	Asn	Arg	Ile	Trp	Ile	Ile	His	Gly	Thr	Ile	Tyr	Trp	
	450					455					460					
atg	tat	acg	gcc	tac	aac	tcg	cgg	acc	ctg	tat	acc	aag	cat	tat	gtg	1440
Met	Tyr	Thr	Ala	Tyr	Asn	Ser	Pro	Thr	Leu	Tyr	Thr	Lys	His	Tyr	Val	
465					470					475					480	
cag	acg	atc	aat	cag	cag	cgg	ctg	gcc	tcc	agc	cgc	tgg	gcc	gcc	tgc	1488
Gln	Thr	Ile	Asn	Gln	Gln	Pro	Leu	Ala	Ser	Ser	Arg	Trp	Ala	Ala	Cys	
			485						490					495		
gcc	att	ggc	ggc	gtg	ctc	gcc	tcc	ttc	atc	caa	att	ctg	gcc	acc	ctc	1536
Ala	Ile	Gly	Gly	Val	Leu	Ala	Ser	Phe	Ile	Gln	Ile	Leu	Ala	Thr	Leu	
			500					505					510			
ttc	gag	tgg	att	ttc	gtt	cgg	cgg	gag	tgg	gcg	ggg	gcg	cag	cat	ctg	1584
Phe	Glu	Trp	Ile	Phe	Val	Pro	Arg	Glu	Trp	Ala	Gly	Ala	Gln	His	Leu	
	515						520						525			
tcg	agg	cgc	atg	ctg	ttc	ctg	gtc	ctc	atc	ttt	ctg	ctt	aac	ctg	gtc	1632
Ser	Arg	Arg	Met	Leu	Phe	Leu	Val	Leu	Ile	Phe	Leu	Leu	Asn	Leu	Val	
	530					535					540					
cct	cgg	gtc	tat	acc	ttc	cag	ata	acc	aag	ctc	gtc	atc	tac	tcg	aag	1680
Pro	Pro	Val	Tyr	Thr	Phe	Gln	Ile	Thr	Lys	Leu	Val	Ile	Tyr	Ser	Lys	
545					550					555					560	
agt	cgg	tac	gct	gtg	tcc	atc	gtg	ggc	ttc	ttc	att	gcc	gtg	gct	acc	1728
Ser	Ala	Tyr	Ala	Val	Ser	Ile	Val	Gly	Phe	Phe	Ile	Ala	Val	Ala	Thr	
			565					570						575		
ctg	gtc	ttt	ttc	cgg	gtc	atg	cgg	ctc	ggg	ggc	ctc	ttc	acg	tcc	tat	1776
Leu	Val	Phe	Phe	Ala	Val	Met	Pro	Leu	Gly	Gly	Leu	Phe	Thr	Ser	Tyr	
		580						585					590			
atg	aac	aag	cgg	agt	cgc	cgg	tat	atc	cgc	agt	cag	acg	ttc	act	gcg	1824
Met	Asn	Lys	Arg	Ser	Arg	Arg	Tyr	Ile	Ala	Ser	Gln	Thr	Phe	Thr	Ala	
		595					600						605			
aac	tat	atc	aag	ctc	aag	ggc	ctc	gac	atg	tgg	atg	agt	tac	ctc	ctg	1872
Asn	Tyr	Ile	Lys	Leu	Lys	Gly	Leu	Asp	Met	Trp	Met	Ser	Tyr	Leu	Leu	
610					615						620					
tgg	ttc	ctg	ggt	ttc	ctc	gcc	aag	ctc	ggt	gag	tcc	tac	ttc	ttc	agc	1920
Trp	Phe	Leu	Val	Phe	Leu	Ala	Lys	Leu	Val	Glu	Ser	Tyr	Phe	Phe	Ser	
625					630					635					640	
acc	ctg	tcg	ctg	cgc	gac	ccc	atc	cgc	aac	ctg	tcc	acg	atg	acg	atg	1968
Thr	Leu	Ser	Leu	Arg	Asp	Pro	Ile	Arg	Asn	Leu	Ser	Thr	Met	Thr	Met	
			645						650					655		
cgc	tgc	gtc	ggc	gag	gtc	tgg	tac	aag	gac	atc	gtg	tgc	cgc	aac	caa	2016
Arg	Cys	Val	Gly	Glu	Val	Trp	Tyr	Lys	Asp	Ile	Val	Cys	Arg	Asn	Gln	
		660						665						670		
gcc	aag	atc	gtc	ctc	ggc	ctc	atg	tat	ctg	gtc	gat	ctg	ctc	ctc	ttc	2064
Ala	Lys	Ile	Val	Leu	Gly	Leu	Met	Tyr	Leu	Val	Asp	Leu	Leu	Leu	Phe	
		675					680					685				
ttc	ctc	gat	acc	tac	atg	tgg	tac	atc	atc	tgc	aac	tgc	atc	ttc	tcc	2112
Phe	Leu	Asp	Thr	Tyr	Met	Trp	Tyr	Ile	Ile	Cys	Asn	Cys	Ile	Phe	Ser	
	690					695					700					
ata	ggc	cgc	agc	ttc	tat	ctg	gga	att	agt	atc	ctc	acc	cgg	tgg	cgg	2160
Ile	Gly	Arg	Ser	Phe	Tyr	Leu	Gly	Ile	Ser	Ile	Leu	Thr	Pro	Trp	Arg	
705					710					715					720	
aac	atc	ttc	acg	cgc	ctg	cgg	aag	cgg	att	tac	agc	aag	atc	ttg	gcg	2208
Asn	Ile	Phe	Thr	Arg	Leu	Pro	Lys	Arg	Ile	Tyr	Ser	Lys	Ile	Leu	Ala	
				725					730					735		
acc	acc	gag	atg	gag	atc	aaa	tac	aaa	ccc	aag	gtc	ctt	atc	agc	cag	2256

-continued

Thr	Thr	Glu	Met	Glu	Ile	Lys	Tyr	Lys	Pro	Lys	Val	Leu	Ile	Ser	Gln	
		740						745					750			
atc	tgg	aat	gcg	att	gtt	atc	tcc	atg	tat	cgc	gag	cat	ctc	ctg	gcc	2304
Ile	Trp	Asn	Ala	Ile	Val	Ile	Ser	Met	Tyr	Arg	Glu	His	Leu	Leu	Ala	
		755						760					765			
atc	gac	cac	gtc	cag	aaa	ctg	ctg	tat	cat	caa	gtg	cca	agc	gag	atc	2352
Ile	Asp	His	Val	Gln	Lys	Leu	Leu	Tyr	His	Gln	Val	Pro	Ser	Glu	Ile	
		770						775					780			
gaa	ggc	aag	cgc	acg	ctg	cgg	gcg	ccg	acg	ttc	ttc	gtg	tcg	caa	gac	2400
Glu	Gly	Lys	Arg	Thr	Leu	Arg	Ala	Pro	Thr	Phe	Phe	Val	Ser	Gln	Asp	
		785				790				795					800	
gac	aat	aat	ttc	gag	act	gag	ttc	ttc	ccg	cgc	aac	tcc	gag	gcc	gag	2448
Asp	Asn	Asn	Phe	Glu	Thr	Glu	Phe	Phe	Pro	Arg	Asn	Ser	Glu	Ala	Glu	
				805						810					815	
cgg	cgc	att	agt	ttc	ttc	gcc	caa	tcc	ctt	gcg	acc	ccg	atg	ccg	gag	2496
Arg	Arg	Ile	Ser	Phe	Phe	Ala	Gln	Ser	Leu	Ala	Thr	Pro	Met	Pro	Glu	
				820					825					830		
ccc	ctc	ccg	gtc	gat	aac	atg	ccc	acg	ttt	acc	gtg	ttc	acc	cct	cat	2544
Pro	Leu	Pro	Val	Asp	Asn	Met	Pro	Thr	Phe	Thr	Val	Phe	Thr	Pro	His	
				835					840					845		
tac	agc	gag	aag	atc	ctg	ctc	agc	ctg	cgc	gag	atc	att	cgc	gag	gac	2592
Tyr	Ser	Glu	Lys	Ile	Leu	Leu	Ser	Leu	Arg	Glu	Ile	Ile	Arg	Glu	Asp	
				850										860		
gac	caa	ttc	tcg	cgc	gtc	acg	ctg	ctg	gag	tat	ctc	aag	cag	ctg	cat	2640
Asp	Gln	Phe	Ser	Arg	Val	Thr	Leu	Leu	Glu	Tyr	Leu	Lys	Gln	Leu	His	
					870										880	
ccg	gtc	gag	tgg	gac	tgc	ttc	ggt	aag	gac	acg	aag	atc	ctc	gcc	gaa	2688
Pro	Val	Glu	Trp	Asp	Cys	Phe	Val	Lys	Asp	Thr	Lys	Ile	Leu	Ala	Glu	
				885						890					895	
gaa	acc	gct	gcc	tac	gag	aac	ggc	gac	gac	tcg	gag	aaa	ttg	tcc	gag	2736
Glu	Thr	Ala	Ala	Tyr	Glu	Asn	Gly	Asp	Asp	Ser	Glu	Lys	Leu	Ser	Glu	
				900						905					910	
gac	ggc	ctc	aag	tcg	aag	atc	gat	gac	ctc	cca	ttc	tac	tgc	atc	ggc	2784
Asp	Gly	Leu	Lys	Ser	Lys	Ile	Asp	Asp	Leu	Pro	Phe	Tyr	Cys	Ile	Gly	
				915					920						925	
ttc	aaa	tcc	gca	gca	ccc	gag	tat	acg	ctc	cgg	acg	cgc	atc	tgg	gcg	2832
Phe	Lys	Ser	Ala	Ala	Pro	Glu	Tyr	Thr	Leu	Arg	Thr	Arg	Ile	Trp	Ala	
				930					935						940	
agc	ctg	cgc	agc	caa	acc	ttg	tac	cgg	acg	gtc	agc	ggc	ttc	atg	aat	2880
Ser	Leu	Arg	Ser	Gln	Thr	Leu	Tyr	Arg	Thr	Val	Ser	Gly	Phe	Met	Asn	
					950										960	
tac	gcc	cgc	gcg	ata	aaa	ctg	ctc	tat	agg	gtc	gag	aac	ccg	gag	ctc	2928
Tyr	Ala	Arg	Ala	Ile	Lys	Leu	Leu	Tyr	Arg	Val	Glu	Asn	Pro	Glu	Leu	
				965						970					975	
gtc	cag	tat	ttt	ggc	ggc	gac	ccc	gag	ggc	ctg	gag	ctg	gcg	ctg	gag	2976
Val	Gln	Tyr	Phe	Gly	Gly	Asp	Pro	Glu	Gly	Leu	Glu	Leu	Ala	Leu	Glu	
				980						985					990	
cgc	atg	gcg	cgc	cgc	aaa	ttc	cgc	ttc	ctc	gtg	agc	atg	cag	cgc	ctg	3024
Arg	Met	Ala	Arg	Arg	Lys	Phe	Arg	Phe	Leu	Val	Ser	Met	Gln	Arg	Leu	
					995				1000						1005	
tcg	aag	ttc	aaa	gac	gat	gag	atg	gag	aat	gct	gaa	ttc	ctc	ctg		3069
Ser	Lys	Phe	Lys	Asp	Asp	Glu	Met	Glu	Asn	Ala	Glu	Phe	Leu	Leu		
						1010								1020		
cgg	gcc	tac	ccg	gac	ctc	cag	atc	gcg	tat	ctc	gac	gaa	gaa	ccg		3114
Arg	Ala	Tyr	Pro	Asp	Leu	Gln	Ile	Ala	Tyr	Leu	Asp	Glu	Glu	Pro		
						1025								1035		
cgc	ctg	aat	gag	gac	gag	gaa	ccg	cgc	gtg	tac	agc	gca	ctg	atc		3159

-continued

Ala 1040	Leu	Asn	Glu	Asp	Glu	Glu	Pro	Arg	Val	Tyr	Ser	Ala	Leu	Ile		
						1045					1050					
gac	gga	cac	tgc	gag	atg	ctg	gag	aac	ggc	agg	cgc	agg	ccg	aag		3204
Asp	Gly	His	Cys	Glu	Met	Leu	Glu	Asn	Gly	Arg	Arg	Arg	Pro	Lys		
	1055					1060					1065					
ttt	cgc	gtc	cag	ttg	tcg	ggc	aac	ccg	atc	ctc	ggc	gat	ggt	aag		3249
Phe	Arg	Val	Gln	Leu	Ser	Gly	Asn	Pro	Ile	Leu	Gly	Asp	Gly	Lys		
	1070					1075					1080					
tcc	gac	aac	cag	aat	cac	gcc	gtc	atc	ttc	cat	cgc	ggc	gag	tat		3294
Ser	Asp	Asn	Gln	Asn	His	Ala	Val	Ile	Phe	His	Arg	Gly	Glu	Tyr		
	1085					1090					1095					
atc	cag	ctg	atc	gac	gcg	aac	cag	gac	aac	tat	ctg	gaa	gag	tgc		3339
Ile	Gln	Leu	Ile	Asp	Ala	Asn	Gln	Asp	Asn	Tyr	Leu	Glu	Glu	Cys		
	1100					1105					1110					
ctg	aag	atc	cgc	tcg	gtc	ctg	gct	gaa	ttc	gaa	gag	atg	aac	gtc		3384
Leu	Lys	Ile	Arg	Ser	Val	Leu	Ala	Glu	Phe	Glu	Glu	Met	Asn	Val		
	1115					1120					1125					
gag	cat	gtt	aat	ccg	tac	gcc	ccg	aat	ttg	aaa	tcc	gag	gac	aac		3429
Glu	His	Val	Asn	Pro	Tyr	Ala	Pro	Asn	Leu	Lys	Ser	Glu	Asp	Asn		
	1130					1135					1140					
aac	acg	aag	aaa	gac	ccg	gtc	gcc	ttc	ctg	ggc	gca	cgc	gag	tac		3474
Asn	Thr	Lys	Lys	Asp	Pro	Val	Ala	Phe	Leu	Gly	Ala	Arg	Glu	Tyr		
	1145					1150					1155					
atc	ttc	agt	gag	aac	tcg	ggc	gtg	ctg	ggc	gac	gtg	gct	gcg	ggc		3519
Ile	Phe	Ser	Glu	Asn	Ser	Gly	Val	Leu	Gly	Asp	Val	Ala	Ala	Gly		
	1160					1165					1170					
aaa	gag	cag	acg	ttt	ggc	acc	ctg	ttc	gcc	cgg	acc	ctc	gcg	cag		3564
Lys	Glu	Gln	Thr	Phe	Gly	Thr	Leu	Phe	Ala	Arg	Thr	Leu	Ala	Gln		
	1175					1180					1185					
atc	ggc	ggg	aag	ctc	cac	tat	ggc	cat	ccc	gac	ttc	ctg	aat	gcg		3609
Ile	Gly	Gly	Lys	Leu	His	Tyr	Gly	His	Pro	Asp	Phe	Leu	Asn	Ala		
	1190					1195					1200					
acg	ttc	atg	ctc	acg	cgc	ggg	ggc	gtg	tcc	aaa	gcc	cag	aag	ggc		3654
Thr	Phe	Met	Leu	Thr	Arg	Gly	Gly	Val	Ser	Lys	Ala	Gln	Lys	Gly		
	1205					1210					1215					
ctg	cac	ctc	aac	gaa	gat	atc	tat	gcg	ggt	atg	aat	gcg	atg	atg		3699
Leu	His	Leu	Asn	Glu	Asp	Ile	Tyr	Ala	Gly	Met	Asn	Ala	Met	Met		
	1220					1225					1230					
cgg	ggt	ggc	aag	atc	aag	cac	tgc	gag	tac	tac	cag	tgc	ggt	aag		3744
Arg	Gly	Gly	Lys	Ile	Lys	His	Cys	Glu	Tyr	Tyr	Gln	Cys	Gly	Lys		
	1235					1240					1245					
ggc	agg	gat	ctt	ggc	ttc	ggc	tcc	atc	ctc	aac	ttc	acc	acc	aag		3789
Gly	Arg	Asp	Leu	Gly	Phe	Gly	Ser	Ile	Leu	Asn	Phe	Thr	Thr	Lys		
	1250					1255					1260					
atc	ggg	gcc	ggc	atg	ggc	gag	caa	atg	ctc	tcg	cgg	gag	tac	ttt		3834
Ile	Gly	Ala	Gly	Met	Gly	Glu	Gln	Met	Leu	Ser	Arg	Glu	Tyr	Phe		
	1265					1270					1275					
tat	ctg	ggt	acc	cag	ctt	ccc	ctt	gac	cgc	ttc	ctc	tcg	ttc	tac		3879
Tyr	Leu	Gly	Thr	Gln	Leu	Pro	Leu	Asp	Arg	Phe	Leu	Ser	Phe	Tyr		
	1280					1285					1290					
tac	ggg	cat	ccg	gga	ttc	cac	atc	aac	aac	ctc	ttc	atc	caa	ctg		3924
Tyr	Gly	His	Pro	Gly	Phe	His	Ile	Asn	Asn	Leu	Phe	Ile	Gln	Leu		
	1295					1300					1305					
agc	ctc	caa	gtg	ttc	atc	ctt	gtg	ctt	ggc	aat	ctg	aat	tcg	ctg		3969
Ser	Leu	Gln	Val	Phe	Ile	Leu	Val	Leu	Gly	Asn	Leu	Asn	Ser	Leu		
	1310					1315					1320					
gcc	cac	gag	gcc	atc	atg	tgt	tcg	tac	aac	aaa	gac	gtg	ccc	gtc		4014

-continued

Ala His	Glu Ala Ile Met Cys	Ser Tyr Asn Lys Asp	Val Pro Val	
1325	1330	1335		
act gac	gtg ctc tac ccc ttc	ggc tgt tac aat atc	gct ccc gcg	4059
Thr Asp	Val Leu Tyr Pro Phe	Gly Cys Tyr Asn Ile	Ala Pro Ala	
1340	1345	1350		
gtg gat	tgg atc cgc cgg tat	acc ctc tcg atc ttc	att gtc ttt	4104
Val Asp	Trp Ile Arg Arg Tyr	Thr Leu Ser Ile Phe	Ile Val Phe	
1355	1360	1365		
ttc atc	tcg ttc atc ccg ctg	gtc gtc cag gag ctc	ata gag cgg	4149
Phe Ile	Ser Phe Ile Pro Leu	Val Val Gln Glu Leu	Ile Glu Arg	
1370	1375	1380		
gga gtg	tgg aag gct ttc cag	cgg ttc gtc cgg cat	ttc atc tcg	4194
Gly Val	Trp Lys Ala Phe Gln	Arg Phe Val Arg His	Phe Ile Ser	
1385	1390	1395		
atg tcc	ccg ttc ttc gag gtg	ttc gtg gcg cag ata	tat tcg agc	4239
Met Ser	Pro Phe Phe Glu Val	Phe Val Ala Gln Ile	Tyr Ser Ser	
1400	1405	1410		
tcc gtg	ttc acg gac ctc acc	gtg ggt ggg gcg agg	tat atc agc	4284
Ser Val	Phe Thr Asp Leu Thr	Val Gly Gly Ala Arg	Tyr Ile Ser	
1415	1420	1425		
acc ggg	cgc ggg ttc gcc acc	tcg cgc atc ccg ttt	tcg atc aag	4329
Thr Gly	Arg Gly Phe Ala Thr	Ser Arg Ile Pro Phe	Ser Ile Lys	
1430	1435	1440		
cgc ttt	gcg gat tcg agt atc	tat atg ggc gct cgg	ctc atg ctc	4374
Arg Phe	Ala Asp Ser Ser Ile	Tyr Met Gly Ala Arg	Leu Met Leu	
1445	1450	1455		
atc ctg	ctt ttc ggt acc gtg	agc cac tgg caa gcg	ccg ctt ctc	4419
Ile Leu	Leu Phe Gly Thr Val	Ser His Trp Gln Ala	Pro Leu Leu	
1460	1465	1470		
tgg ttc	tgg gcc tcg ctc tcc	gcg ttg atg ttc tcc	ccg ttc atc	4464
Trp Phe	Trp Ala Ser Leu Ser	Ala Leu Met Phe Ser	Pro Phe Ile	
1475	1480	1485		
ttt aac	ccg cat cag ttc gcg	tgg gaa gat ttc ttt	ctc gac tat	4509
Phe Asn	Pro His Gln Phe Ala	Trp Glu Asp Phe Phe	Leu Asp Tyr	
1490	1495	1500		
cgc gat	ttc atc cgc tgg ctg	tcc cgc gga aac acc	aag tgg cat	4554
Arg Asp	Phe Ile Arg Trp Leu	Ser Arg Gly Asn Thr	Lys Trp His	
1505	1510	1515		
cgg aac	agc tgg atc ggc tac	gtt cgc ctc tcg cgg	tcg cgc ata	4599
Arg Asn	Ser Trp Ile Gly Tyr	Val Arg Leu Ser Arg	Ser Arg Ile	
1520	1525	1530		
acg ggc	ttc aag cgg aaa ctc	acc ggc gac gtg agc	gag aaa gcc	4644
Thr Gly	Phe Lys Arg Lys Leu	Thr Gly Asp Val Ser	Glu Lys Ala	
1535	1540	1545		
gct ggc	gat gcc tcc agg gct	cat cgc tcg aat gtg	ctg ttc gcg	4689
Ala Gly	Asp Ala Ser Arg Ala	His Arg Ser Asn Val	Leu Phe Ala	
1550	1555	1560		
gac ttc	ctg ccg acc ctg ata	tac act gcg ggc ctc	tac gtc gcc	4734
Asp Phe	Leu Pro Thr Leu Ile	Tyr Thr Ala Gly Leu	Tyr Val Ala	
1565	1570	1575		
tat acc	ttc atc aac gcc cag	acg ggc gtc acc agc	tat ccc tac	4779
Tyr Thr	Phe Ile Asn Ala Gln	Thr Gly Val Thr Ser	Tyr Pro Tyr	
1580	1585	1590		
gag atc	aat ggc tcg acg gac	ccg caa cca gtt aac	tcc acg ctg	4824
Glu Ile	Asn Gly Ser Thr Asp	Pro Gln Pro Val Asn	Ser Thr Leu	
1595	1600	1605		
agg ctc	atc atc tgc gcc ctg	gca cct gtc gtc atc	gac atg gcc	4869

-continued

Arg 1610	Leu	Ile	Ile	Cys	Ala	Leu 1615	Ala	Pro	Val	Val	Ile 1620	Asp	Met	Gly	
tgc	ctg	ggc	gtg	tgc	ctc	gcc	atg	gcg	tgc	tgc	gcc	ggc	ccg	atg	4914
Cys 1625	Leu	Gly	Val	Cys	Leu	Ala 1630	Met	Ala	Cys	Cys	Ala 1635	Gly	Pro	Met	
ctg	ggc	ctg	tgc	tgc	aag	aaa	acc	ggc	gca	gtc	atc	gcc	ggc	gtc	4959
Leu 1640	Gly	Leu	Cys	Cys	Lys	Lys 1645	Thr	Gly	Ala	Val	Ile 1650	Ala	Gly	Val	
gcg	cac	ggc	gtc	gcc	gtc	atc	gtc	cat	atc	atc	ttc	ttc	atc	gtg	5004
Ala 1655	His	Gly	Val	Ala	Val	Ile 1660	Val	His	Ile	Ile	Phe 1665	Phe	Ile	Val	
atg	tgg	gtg	act	gag	ggc	ttc	aat	ttc	gca	cgc	ctg	atg	ctt	ggc	5049
Met 1670	Trp	Val	Thr	Glu	Gly	Phe 1675	Asn	Phe	Ala	Arg	Leu 1680	Met	Leu	Gly	
atc	gcg	acc	atg	atc	tat	gtg	cag	agg	ctg	ctt	ttc	aag	ttc	ctg	5094
Ile 1685	Ala	Thr	Met	Ile	Tyr	Val 1690	Gln	Arg	Leu	Leu	Phe 1695	Lys	Phe	Leu	
acg	ctg	tgc	ttc	ctc	acc	cgg	gag	ttc	aag	aac	gac	aaa	gcc	aac	5139
Thr 1700	Leu	Cys	Phe	Leu	Thr	Arg 1705	Glu	Phe	Lys	Asn	Asp 1710	Lys	Ala	Asn	
acc	gcg	ttt	tgg	acg	ggc	aag	tgg	tat	aac	acc	ggc	atg	ggc	tgg	5184
Thr 1715	Ala	Phe	Trp	Thr	Gly	Lys 1720	Trp	Tyr	Asn	Thr	Gly 1725	Met	Gly	Trp	
atg	gcc	ttc	acg	cag	ccg	agc	cgc	gag	ttc	gtg	gcc	aag	atc	atc	5229
Met 1730	Ala	Phe	Thr	Gln	Pro	Ser 1735	Arg	Glu	Phe	Val	Ala 1740	Lys	Ile	Ile	
gag	atg	agc	gag	ttc	gcg	ggc	gat	ttc	gtc	ctt	gcc	cac	ata	atc	5274
Glu 1745	Met	Ser	Glu	Phe	Ala	Gly 1750	Asp	Phe	Val	Leu	Ala 1755	His	Ile	Ile	
ctg	ttc	tgc	cag	ctg	ccg	ctc	ctg	ttc	att	ccg	ttg	gtc	gac	cgc	5319
Leu 1760	Phe	Cys	Gln	Leu	Pro	Leu 1765	Leu	Phe	Ile	Pro	Leu 1770	Val	Asp	Arg	
tgg	cat	agc	atg	atg	ctg	ttc	tgg	ctg	aag	ccg	tcc	cgc	ctc	att	5364
Trp 1775	His	Ser	Met	Met	Leu	Phe 1780	Trp	Leu	Lys	Pro	Ser 1785	Arg	Leu	Ile	
cgg	cca	cct	atc	tac	tcg	ctc	aaa	caa	gcg	cgc	ctg	cgc	aag	cgg	5409
Arg 1790	Pro	Pro	Ile	Tyr	Ser	Leu 1795	Lys	Gln	Ala	Arg	Leu 1800	Arg	Lys	Arg	
atg	gtc	cgc	aaa	tac	tgt	gtc	ctg	tat	ttc	gca	gtg	ctt	atc	ctg	5454
Met 1805	Val	Arg	Lys	Tyr	Cys	Val 1810	Leu	Tyr	Phe	Ala	Val 1815	Leu	Ile	Leu	
ttc	atc	ggt	atc	atc	gtc	gcg	cct	gcg	gtc	gca	tcg	ggg	cag	atc	5499
Phe 1820	Ile	Val	Ile	Ile	Val	Ala 1825	Pro	Ala	Val	Ala	Ser 1830	Gly	Gln	Ile	
gcg	gtg	gac	cag	ttc	gcc	aac	att	ggc	ggc	agc	ggc	tcc	ata	gca	5544
Ala 1835	Val	Asp	Gln	Phe	Ala	Asn 1840	Ile	Gly	Gly	Ser	Gly 1845	Ser	Ile	Ala	
gac	ggc	ttg	ttc	caa	ccc	cgg	aat	gtg	agc	aac	aac	gac	acc	ggg	5589
Asp 1850	Gly	Leu	Phe	Gln	Pro	Arg 1855	Asn	Val	Ser	Asn	Asn 1860	Asp	Thr	Gly	
aat	cat	cgc	cca	aag	acc	tac	acc	tgg	tcc	tac	ctc	tcg	acc	cgc	5634
Asn 1865	His	Arg	Pro	Lys	Thr	Tyr 1870	Thr	Trp	Ser	Tyr	Leu 1875	Ser	Thr	Arg	
ttc	acc	ggc	tcg	acc	acc	ccc	tac	tcc	acc	aat	ccc	ttc	agg	gtg	5679
Phe 1880	Thr	Gly	Ser	Thr	Thr	Pro 1885	Tyr	Ser	Thr	Asn	Pro 1890	Phe	Arg	Val	

-continued

```

<210> SEQ ID NO 10
<211> LENGTH: 1893
<212> TYPE: PRT
<213> ORGANISM: Artificial sequence
<220> FEATURE:
<223> OTHER INFORMATION: Synthetic Construct

<400> SEQUENCE: 10

Met Ser Tyr Asn Asp Asn Asn Asn His Tyr Tyr Asp Pro Asn Gln Gln
1      5      10      15

Gly Gly Met Pro Pro His Gln Gly Gly Glu Gly Tyr Tyr Gln Gln Gln
20      25      30

Tyr Asp Asp Met Gly Gln Gln Pro His Gln Gln Asp Tyr Tyr Asp Pro
35      40      45

Asn Ala Gln Tyr Gln Gln Gln Pro Tyr Asp Met Asp Gly Tyr Gln Asp
50      55      60

Gln Ala Asn Tyr Gly Gly Gln Pro Met Asn Ala Gln Gly Tyr Asn Ala
65      70      75      80

Asp Pro Glu Ala Phe Ser Asp Phe Ser Tyr Gly Gly Gln Thr Pro Gly
85      90      95

Thr Pro Gly Tyr Asp Gln Tyr Gly Thr Gln Tyr Thr Pro Ser Gln Met
100     105     110

Ser Tyr Gly Gly Asp Pro Arg Ser Ser Gly Ala Ser Thr Pro Ile Tyr
115     120     125

Gly Gly Gln Gly Gln Gly Tyr Asp Pro Thr Gln Phe Asn Met Ser Ser
130     135     140

Asn Leu Pro Tyr Pro Ala Trp Ser Ala Asp Pro Gln Ala Pro Ile Lys
145     150     155     160

Ile Glu His Ile Glu Asp Ile Phe Ile Asp Leu Thr Asn Lys Phe Gly
165     170     175

Phe Gln Arg Asp Ser Met Arg Asn Met Phe Asp Tyr Phe Met Thr Leu
180     185     190

Leu Asp Ser Arg Ser Ser Arg Met Ser Pro Ala Gln Ala Leu Leu Ser
195     200     205

Leu His Ala Asp Tyr Ile Gly Gly Asp Asn Ala Asn Tyr Arg Lys Trp
210     215     220

Tyr Phe Ser Ser Gln Gln Asp Leu Asp Asp Ser Leu Gly Phe Ala Asn
225     230     235     240

Met Thr Leu Gly Lys Ile Gly Arg Lys Ala Arg Lys Ala Ser Lys Lys
245     250     255

Ser Lys Lys Ala Arg Lys Ala Ala Glu Glu His Gly Gln Asp Val Asp
260     265     270

Ala Asn Glu Leu Glu Gly Asp Tyr Ser Leu Glu Ala Ala Glu Ile Arg
275     280     285

Trp Lys Ala Lys Met Asn Ser Leu Thr Pro Glu Glu Arg Val Arg Asp
290     295     300

Leu Ala Leu Tyr Leu Leu Ile Trp Gly Glu Ala Asn Gln Val Arg Phe
305     310     315     320

Thr Pro Glu Cys Leu Cys Tyr Ile Tyr Lys Ser Ala Thr Asp Tyr Leu
325     330     335

Asn Ser Pro Leu Cys Gln Gln Arg Gln Glu Pro Val Pro Glu Gly Asp
340     345     350

Tyr Leu Asn Arg Val Ile Thr Pro Leu Tyr Arg Phe Ile Arg Ser Gln

```

-continued

355					360					365					
Val	Tyr	Glu	Ile	Tyr	Asp	Gly	Arg	Phe	Val	Lys	Arg	Glu	Lys	Asp	His
370						375					380				
Asn	Lys	Val	Ile	Gly	Tyr	Asp	Asp	Val	Asn	Gln	Leu	Phe	Trp	Tyr	Pro
385					390					395					400
Glu	Gly	Ile	Ser	Arg	Ile	Ile	Phe	Glu	Asp	Gly	Thr	Arg	Leu	Val	Asp
				405					410						415
Ile	Pro	Gln	Glu	Glu	Arg	Phe	Leu	Lys	Leu	Gly	Glu	Val	Glu	Trp	Lys
				420				425							430
Asn	Val	Phe	Phe	Lys	Thr	Tyr	Lys	Glu	Ile	Arg	Thr	Trp	Leu	His	Phe
				435				440							445
Val	Thr	Asn	Phe	Asn	Arg	Ile	Trp	Ile	Ile	His	Gly	Thr	Ile	Tyr	Trp
450						455					460				
Met	Tyr	Thr	Ala	Tyr	Asn	Ser	Pro	Thr	Leu	Tyr	Thr	Lys	His	Tyr	Val
465					470						475				480
Gln	Thr	Ile	Asn	Gln	Gln	Pro	Leu	Ala	Ser	Ser	Arg	Trp	Ala	Ala	Cys
				485					490						495
Ala	Ile	Gly	Gly	Val	Leu	Ala	Ser	Phe	Ile	Gln	Ile	Leu	Ala	Thr	Leu
				500					505						510
Phe	Glu	Trp	Ile	Phe	Val	Pro	Arg	Glu	Trp	Ala	Gly	Ala	Gln	His	Leu
				515					520						525
Ser	Arg	Arg	Met	Leu	Phe	Leu	Val	Leu	Ile	Phe	Leu	Leu	Asn	Leu	Val
530						535					540				
Pro	Pro	Val	Tyr	Thr	Phe	Gln	Ile	Thr	Lys	Leu	Val	Ile	Tyr	Ser	Lys
545						550					555				560
Ser	Ala	Tyr	Ala	Val	Ser	Ile	Val	Gly	Phe	Phe	Ile	Ala	Val	Ala	Thr
				565					570						575
Leu	Val	Phe	Phe	Ala	Val	Met	Pro	Leu	Gly	Gly	Leu	Phe	Thr	Ser	Tyr
				580					585						590
Met	Asn	Lys	Arg	Ser	Arg	Arg	Tyr	Ile	Ala	Ser	Gln	Thr	Phe	Thr	Ala
				595					600						605
Asn	Tyr	Ile	Lys	Leu	Lys	Gly	Leu	Asp	Met	Trp	Met	Ser	Tyr	Leu	Leu
610						615					620				
Trp	Phe	Leu	Val	Phe	Leu	Ala	Lys	Leu	Val	Glu	Ser	Tyr	Phe	Phe	Ser
625					630						635				640
Thr	Leu	Ser	Leu	Arg	Asp	Pro	Ile	Arg	Asn	Leu	Ser	Thr	Met	Thr	Met
				645					650						655
Arg	Cys	Val	Gly	Glu	Val	Trp	Tyr	Lys	Asp	Ile	Val	Cys	Arg	Asn	Gln
				660					665						670
Ala	Lys	Ile	Val	Leu	Gly	Leu	Met	Tyr	Leu	Val	Asp	Leu	Leu	Leu	Phe
				675					680						685
Phe	Leu	Asp	Thr	Tyr	Met	Trp	Tyr	Ile	Ile	Cys	Asn	Cys	Ile	Phe	Ser
690						695					700				
Ile	Gly	Arg	Ser	Phe	Tyr	Leu	Gly	Ile	Ser	Ile	Leu	Thr	Pro	Trp	Arg
705					710						715				720
Asn	Ile	Phe	Thr	Arg	Leu	Pro	Lys	Arg	Ile	Tyr	Ser	Lys	Ile	Leu	Ala
				725					730						735
Thr	Thr	Glu	Met	Glu	Ile	Lys	Tyr	Lys	Pro	Lys	Val	Leu	Ile	Ser	Gln
				740					745						750
Ile	Trp	Asn	Ala	Ile	Val	Ile	Ser	Met	Tyr	Arg	Glu	His	Leu	Leu	Ala
				755					760						765

-continued

Ile Asp His Val Gln Lys Leu Leu Tyr His Gln Val Pro Ser Glu Ile
 770 775 780
 Glu Gly Lys Arg Thr Leu Arg Ala Pro Thr Phe Phe Val Ser Gln Asp
 785 790 795 800
 Asp Asn Asn Phe Glu Thr Glu Phe Phe Pro Arg Asn Ser Glu Ala Glu
 805 810 815
 Arg Arg Ile Ser Phe Phe Ala Gln Ser Leu Ala Thr Pro Met Pro Glu
 820 825 830
 Pro Leu Pro Val Asp Asn Met Pro Thr Phe Thr Val Phe Thr Pro His
 835 840 845
 Tyr Ser Glu Lys Ile Leu Leu Ser Leu Arg Glu Ile Ile Arg Glu Asp
 850 855 860
 Asp Gln Phe Ser Arg Val Thr Leu Leu Glu Tyr Leu Lys Gln Leu His
 865 870 875 880
 Pro Val Glu Trp Asp Cys Phe Val Lys Asp Thr Lys Ile Leu Ala Glu
 885 890 895
 Glu Thr Ala Ala Tyr Glu Asn Gly Asp Asp Ser Glu Lys Leu Ser Glu
 900 905 910
 Asp Gly Leu Lys Ser Lys Ile Asp Asp Leu Pro Phe Tyr Cys Ile Gly
 915 920 925
 Phe Lys Ser Ala Ala Pro Glu Tyr Thr Leu Arg Thr Arg Ile Trp Ala
 930 935 940
 Ser Leu Arg Ser Gln Thr Leu Tyr Arg Thr Val Ser Gly Phe Met Asn
 945 950 955 960
 Tyr Ala Arg Ala Ile Lys Leu Leu Tyr Arg Val Glu Asn Pro Glu Leu
 965 970 975
 Val Gln Tyr Phe Gly Gly Asp Pro Glu Gly Leu Glu Leu Ala Leu Glu
 980 985 990
 Arg Met Ala Arg Arg Lys Phe Arg Phe Leu Val Ser Met Gln Arg Leu
 995 1000 1005
 Ser Lys Phe Lys Asp Asp Glu Met Glu Asn Ala Glu Phe Leu Leu
 1010 1015 1020
 Arg Ala Tyr Pro Asp Leu Gln Ile Ala Tyr Leu Asp Glu Glu Pro
 1025 1030 1035
 Ala Leu Asn Glu Asp Glu Glu Pro Arg Val Tyr Ser Ala Leu Ile
 1040 1045 1050
 Asp Gly His Cys Glu Met Leu Glu Asn Gly Arg Arg Arg Pro Lys
 1055 1060 1065
 Phe Arg Val Gln Leu Ser Gly Asn Pro Ile Leu Gly Asp Gly Lys
 1070 1075 1080
 Ser Asp Asn Gln Asn His Ala Val Ile Phe His Arg Gly Glu Tyr
 1085 1090 1095
 Ile Gln Leu Ile Asp Ala Asn Gln Asp Asn Tyr Leu Glu Glu Cys
 1100 1105 1110
 Leu Lys Ile Arg Ser Val Leu Ala Glu Phe Glu Glu Met Asn Val
 1115 1120 1125
 Glu His Val Asn Pro Tyr Ala Pro Asn Leu Lys Ser Glu Asp Asn
 1130 1135 1140
 Asn Thr Lys Lys Asp Pro Val Ala Phe Leu Gly Ala Arg Glu Tyr
 1145 1150 1155

-continued

Ile	Phe	Ser	Glu	Asn	Ser	Gly	Val	Leu	Gly	Asp	Val	Ala	Ala	Gly
1160						1165					1170			
Lys	Glu	Gln	Thr	Phe	Gly	Thr	Leu	Phe	Ala	Arg	Thr	Leu	Ala	Gln
1175						1180					1185			
Ile	Gly	Gly	Lys	Leu	His	Tyr	Gly	His	Pro	Asp	Phe	Leu	Asn	Ala
1190						1195					1200			
Thr	Phe	Met	Leu	Thr	Arg	Gly	Gly	Val	Ser	Lys	Ala	Gln	Lys	Gly
1205						1210					1215			
Leu	His	Leu	Asn	Glu	Asp	Ile	Tyr	Ala	Gly	Met	Asn	Ala	Met	Met
1220						1225					1230			
Arg	Gly	Gly	Lys	Ile	Lys	His	Cys	Glu	Tyr	Tyr	Gln	Cys	Gly	Lys
1235						1240					1245			
Gly	Arg	Asp	Leu	Gly	Phe	Gly	Ser	Ile	Leu	Asn	Phe	Thr	Thr	Lys
1250						1255					1260			
Ile	Gly	Ala	Gly	Met	Gly	Glu	Gln	Met	Leu	Ser	Arg	Glu	Tyr	Phe
1265						1270					1275			
Tyr	Leu	Gly	Thr	Gln	Leu	Pro	Leu	Asp	Arg	Phe	Leu	Ser	Phe	Tyr
1280						1285					1290			
Tyr	Gly	His	Pro	Gly	Phe	His	Ile	Asn	Asn	Leu	Phe	Ile	Gln	Leu
1295						1300					1305			
Ser	Leu	Gln	Val	Phe	Ile	Leu	Val	Leu	Gly	Asn	Leu	Asn	Ser	Leu
1310						1315					1320			
Ala	His	Glu	Ala	Ile	Met	Cys	Ser	Tyr	Asn	Lys	Asp	Val	Pro	Val
1325						1330					1335			
Thr	Asp	Val	Leu	Tyr	Pro	Phe	Gly	Cys	Tyr	Asn	Ile	Ala	Pro	Ala
1340						1345					1350			
Val	Asp	Trp	Ile	Arg	Arg	Tyr	Thr	Leu	Ser	Ile	Phe	Ile	Val	Phe
1355						1360					1365			
Phe	Ile	Ser	Phe	Ile	Pro	Leu	Val	Val	Gln	Glu	Leu	Ile	Glu	Arg
1370						1375					1380			
Gly	Val	Trp	Lys	Ala	Phe	Gln	Arg	Phe	Val	Arg	His	Phe	Ile	Ser
1385						1390					1395			
Met	Ser	Pro	Phe	Phe	Glu	Val	Phe	Val	Ala	Gln	Ile	Tyr	Ser	Ser
1400						1405					1410			
Ser	Val	Phe	Thr	Asp	Leu	Thr	Val	Gly	Gly	Ala	Arg	Tyr	Ile	Ser
1415						1420					1425			
Thr	Gly	Arg	Gly	Phe	Ala	Thr	Ser	Arg	Ile	Pro	Phe	Ser	Ile	Lys
1430						1435					1440			
Arg	Phe	Ala	Asp	Ser	Ser	Ile	Tyr	Met	Gly	Ala	Arg	Leu	Met	Leu
1445						1450					1455			
Ile	Leu	Leu	Phe	Gly	Thr	Val	Ser	His	Trp	Gln	Ala	Pro	Leu	Leu
1460						1465					1470			
Trp	Phe	Trp	Ala	Ser	Leu	Ser	Ala	Leu	Met	Phe	Ser	Pro	Phe	Ile
1475						1480					1485			
Phe	Asn	Pro	His	Gln	Phe	Ala	Trp	Glu	Asp	Phe	Phe	Leu	Asp	Tyr
1490						1495					1500			
Arg	Asp	Phe	Ile	Arg	Trp	Leu	Ser	Arg	Gly	Asn	Thr	Lys	Trp	His
1505						1510					1515			
Arg	Asn	Ser	Trp	Ile	Gly	Tyr	Val	Arg	Leu	Ser	Arg	Ser	Arg	Ile
1520						1525					1530			
Thr	Gly	Phe	Lys	Arg	Lys	Leu	Thr	Gly	Asp	Val	Ser	Glu	Lys	Ala

-continued

1535	1540	1545
Ala Gly Asp Ala Ser Arg Ala His Arg Ser Asn Val Leu Phe Ala 1550 1555 1560		
Asp Phe Leu Pro Thr Leu Ile Tyr Thr Ala Gly Leu Tyr Val Ala 1565 1570 1575		
Tyr Thr Phe Ile Asn Ala Gln Thr Gly Val Thr Ser Tyr Pro Tyr 1580 1585 1590		
Glu Ile Asn Gly Ser Thr Asp Pro Gln Pro Val Asn Ser Thr Leu 1595 1600 1605		
Arg Leu Ile Ile Cys Ala Leu Ala Pro Val Val Ile Asp Met Gly 1610 1615 1620		
Cys Leu Gly Val Cys Leu Ala Met Ala Cys Cys Ala Gly Pro Met 1625 1630 1635		
Leu Gly Leu Cys Cys Lys Lys Thr Gly Ala Val Ile Ala Gly Val 1640 1645 1650		
Ala His Gly Val Ala Val Ile Val His Ile Ile Phe Phe Ile Val 1655 1660 1665		
Met Trp Val Thr Glu Gly Phe Asn Phe Ala Arg Leu Met Leu Gly 1670 1675 1680		
Ile Ala Thr Met Ile Tyr Val Gln Arg Leu Leu Phe Lys Phe Leu 1685 1690 1695		
Thr Leu Cys Phe Leu Thr Arg Glu Phe Lys Asn Asp Lys Ala Asn 1700 1705 1710		
Thr Ala Phe Trp Thr Gly Lys Trp Tyr Asn Thr Gly Met Gly Trp 1715 1720 1725		
Met Ala Phe Thr Gln Pro Ser Arg Glu Phe Val Ala Lys Ile Ile 1730 1735 1740		
Glu Met Ser Glu Phe Ala Gly Asp Phe Val Leu Ala His Ile Ile 1745 1750 1755		
Leu Phe Cys Gln Leu Pro Leu Leu Phe Ile Pro Leu Val Asp Arg 1760 1765 1770		
Trp His Ser Met Met Leu Phe Trp Leu Lys Pro Ser Arg Leu Ile 1775 1780 1785		
Arg Pro Pro Ile Tyr Ser Leu Lys Gln Ala Arg Leu Arg Lys Arg 1790 1795 1800		
Met Val Arg Lys Tyr Cys Val Leu Tyr Phe Ala Val Leu Ile Leu 1805 1810 1815		
Phe Ile Val Ile Ile Val Ala Pro Ala Val Ala Ser Gly Gln Ile 1820 1825 1830		
Ala Val Asp Gln Phe Ala Asn Ile Gly Gly Ser Gly Ser Ile Ala 1835 1840 1845		
Asp Gly Leu Phe Gln Pro Arg Asn Val Ser Asn Asn Asp Thr Gly 1850 1855 1860		
Asn His Arg Pro Lys Thr Tyr Thr Trp Ser Tyr Leu Ser Thr Arg 1865 1870 1875		
Phe Thr Gly Ser Thr Thr Pro Tyr Ser Thr Asn Pro Phe Arg Val 1880 1885 1890		

<210> SEQ ID NO 11

<211> LENGTH: 171

<212> TYPE: DNA

<213> ORGANISM: Artificial sequence

-continued

```

<220> FEATURE:
<223> OTHER INFORMATION: Synthetic construct
<220> FEATURE:
<221> NAME/KEY: CDS
<222> LOCATION: (1)..(171)
<223> OTHER INFORMATION: Zea mays (corn) portion of 1,3-D-glucan
      synthase (codon optimized)

<400> SEQUENCE: 11

ttc aac tgt acc ctc cgc ggc ggc aat gtt acc cat cac gaa tat atc      48
Phe Asn Cys Thr Leu Arg Gly Gly Asn Val Thr His His Glu Tyr Ile
1           5                10                15

caa gtc ggc aaa gga cgc gac gtc ggc ctg aat caa gtg tcg atg ttc      96
Gln Val Gly Lys Gly Arg Asp Val Gly Leu Asn Gln Val Ser Met Phe
          20                25                30

gag cgc aaa gtc gcc tcc ggt aac ggc gag cag acg ctg agc cgc gac     144
Glu Ala Lys Val Ala Ser Gly Asn Gly Glu Gln Thr Leu Ser Arg Asp
          35                40                45

gtg tac cgg ctc ggg cat cgg ctg gat                                  171
Val Tyr Arg Leu Gly His Arg Leu Asp
          50                55

<210> SEQ ID NO 12
<211> LENGTH: 57
<212> TYPE: PRT
<213> ORGANISM: Artificial sequence
<220> FEATURE:
<223> OTHER INFORMATION: Synthetic Construct

<400> SEQUENCE: 12

Phe Asn Cys Thr Leu Arg Gly Gly Asn Val Thr His His Glu Tyr Ile
1           5                10                15

Gln Val Gly Lys Gly Arg Asp Val Gly Leu Asn Gln Val Ser Met Phe
          20                25                30

Glu Ala Lys Val Ala Ser Gly Asn Gly Glu Gln Thr Leu Ser Arg Asp
          35                40                45

Val Tyr Arg Leu Gly His Arg Leu Asp
          50                55

<210> SEQ ID NO 13
<211> LENGTH: 450
<212> TYPE: DNA
<213> ORGANISM: Artificial sequence
<220> FEATURE:
<223> OTHER INFORMATION: Synthetic construct
<220> FEATURE:
<221> NAME/KEY: CDS
<222> LOCATION: (1)..(450)
<223> OTHER INFORMATION: Zea mays (corn): portion of 1,3-D-glucan
      synthase (codon optimized)

<400> SEQUENCE: 13

cag ctg aag cgc ctg cat ctc ctc ctc acc gtc aaa gac agc gcc acc      48
Gln Leu Lys Arg Leu His Leu Leu Leu Thr Val Lys Asp Ser Ala Thr
1           5                10                15

aac atc ccg aag aat ctt gag gcc cgg cgg cgc ctg cag ttc ttc acg      96
Asn Ile Pro Lys Asn Leu Glu Ala Arg Arg Arg Leu Gln Phe Phe Thr
          20                25                30

aac agc ctg ttc atg gat atc ccg caa gcg aag ccc gtg tcc gag atg     144
Asn Ser Leu Phe Met Asp Ile Pro Gln Ala Lys Pro Val Ser Glu Met
          35                40                45

atc ccg ttt tcg gtg ttc acc ccg tac tac tcg gag act gtt ctc tat     192

```

-continued

```

Ile Pro Phe Ser Val Phe Thr Pro Tyr Tyr Ser Glu Thr Val Leu Tyr
 50          55          60

tcc atg tcc gag ctg tgc gtc gag aat gag gac ggc atc agt att ctg      240
Ser Met Ser Glu Leu Cys Val Glu Asn Glu Asp Gly Ile Ser Ile Leu
65          70          75          80

ttc tac ctc caa aag atc tat ccc gac gag tgg gca aac ttc ctg gag      288
Phe Tyr Leu Gln Lys Ile Tyr Pro Asp Glu Trp Ala Asn Phe Leu Glu
85          90          95

cgc atc ggg tgc ggc gag tcg agc gaa gat gac ttc aaa gaa tcg ccg      336
Arg Ile Gly Cys Gly Glu Ser Ser Glu Asp Asp Phe Lys Glu Ser Pro
100         105         110

tcc gac acg atg gaa ttg cgg ttc tgg gtg agc tac cgc ggt cag acc      384
Ser Asp Thr Met Glu Leu Arg Phe Trp Val Ser Tyr Arg Gly Gln Thr
115         120         125

ctc ggc cgc acc gtc cgg ggc atg atg tat tac cgc agg gcg ctg atg      432
Leu Gly Arg Thr Val Arg Gly Met Met Tyr Tyr Arg Arg Ala Leu Met
130         135         140

ctc cag tcg tat ctg gag      450
Leu Gln Ser Tyr Leu Glu
145         150

```

```

<210> SEQ ID NO 14
<211> LENGTH: 150
<212> TYPE: PRT
<213> ORGANISM: Artificial sequence
<220> FEATURE:
<223> OTHER INFORMATION: Synthetic Construct

```

<400> SEQUENCE: 14

```

Gln Leu Lys Arg Leu His Leu Leu Leu Thr Val Lys Asp Ser Ala Thr
 1          5          10          15

Asn Ile Pro Lys Asn Leu Glu Ala Arg Arg Arg Leu Gln Phe Phe Thr
20         25         30

Asn Ser Leu Phe Met Asp Ile Pro Gln Ala Lys Pro Val Ser Glu Met
35         40         45

Ile Pro Phe Ser Val Phe Thr Pro Tyr Tyr Ser Glu Thr Val Leu Tyr
50          55          60

Ser Met Ser Glu Leu Cys Val Glu Asn Glu Asp Gly Ile Ser Ile Leu
65          70          75          80

Phe Tyr Leu Gln Lys Ile Tyr Pro Asp Glu Trp Ala Asn Phe Leu Glu
85          90          95

Arg Ile Gly Cys Gly Glu Ser Ser Glu Asp Asp Phe Lys Glu Ser Pro
100         105         110

Ser Asp Thr Met Glu Leu Arg Phe Trp Val Ser Tyr Arg Gly Gln Thr
115         120         125

Leu Gly Arg Thr Val Arg Gly Met Met Tyr Tyr Arg Arg Ala Leu Met
130         135         140

Leu Gln Ser Tyr Leu Glu
145         150

```

```

<210> SEQ ID NO 15
<211> LENGTH: 405
<212> TYPE: DNA
<213> ORGANISM: Artificial sequence
<220> FEATURE:
<223> OTHER INFORMATION: Synthetic construct
<220> FEATURE:
<221> NAME/KEY: CDS

```

-continued

<222> LOCATION: (1)..(405)
 <223> OTHER INFORMATION: Oryza sativa (rice) portion of 1,3-beta-D-glucan synthase (codon optimized)

<400> SEQUENCE: 15

```

gag aac tat cgc ctc tac tcg cgc tcc cat ttc gtc aaa gca ctc gaa      48
Glu Asn Tyr Arg Leu Tyr Ser Arg Ser His Phe Val Lys Ala Leu Glu
1           5           10           15

gtt gcc ctg ctg ctc atc atc tat att gcg tac ggc tat acc cgg ggg      96
Val Ala Leu Leu Leu Ile Ile Tyr Ile Ala Tyr Gly Tyr Thr Arg Gly
           20           25           30

ggc tcc tcc agc ttc atc ctg ttg acc att agt tcg tgg ttc ctc gtc     144
Gly Ser Ser Ser Phe Ile Leu Leu Thr Ile Ser Ser Trp Phe Leu Val
           35           40           45

gtg tcg tgg ctg ttc gct ccc tac ata ttc aac ccg agc ggc ttt gag     192
Val Ser Trp Leu Phe Ala Pro Tyr Ile Phe Asn Pro Ser Gly Phe Glu
           50           55           60

tgg cag aaa acc gtc gag gac ttc gac gat tgg acg aat tgg ctc ctg     240
Trp Gln Lys Thr Val Glu Asp Phe Asp Asp Trp Thr Asn Trp Leu Leu
65           70           75           80

tac aag ggc gga gtg ggc gtg aag ggt gag aat agc tgg gag tcc tgg     288
Tyr Lys Gly Gly Val Gly Val Lys Gly Glu Asn Ser Trp Glu Ser Trp
           85           90           95

tgg gac gag gaa cag gcg cat atc caa acg ctg agg ggt cgg atc ctt     336
Trp Asp Glu Glu Gln Ala His Ile Gln Thr Leu Arg Gly Arg Ile Leu
           100          105          110

gag act atc ctg tcg ctg cgc ttc ctc atc ttc cag tac ggc atc gtc     384
Glu Thr Ile Leu Ser Leu Arg Phe Leu Ile Phe Gln Tyr Gly Ile Val
           115          120          125

tat aag ctc aag atc gcc cac      405
Tyr Lys Leu Lys Ile Ala His
           130          135
    
```

<210> SEQ ID NO 16
 <211> LENGTH: 135
 <212> TYPE: PRT
 <213> ORGANISM: Artificial sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Synthetic Construct

<400> SEQUENCE: 16

```

Glu Asn Tyr Arg Leu Tyr Ser Arg Ser His Phe Val Lys Ala Leu Glu
1           5           10           15

Val Ala Leu Leu Leu Ile Ile Tyr Ile Ala Tyr Gly Tyr Thr Arg Gly
           20           25           30

Gly Ser Ser Ser Phe Ile Leu Leu Thr Ile Ser Ser Trp Phe Leu Val
           35           40           45

Val Ser Trp Leu Phe Ala Pro Tyr Ile Phe Asn Pro Ser Gly Phe Glu
           50           55           60

Trp Gln Lys Thr Val Glu Asp Phe Asp Asp Trp Thr Asn Trp Leu Leu
65           70           75           80

Tyr Lys Gly Gly Val Gly Val Lys Gly Glu Asn Ser Trp Glu Ser Trp
           85           90           95

Trp Asp Glu Glu Gln Ala His Ile Gln Thr Leu Arg Gly Arg Ile Leu
           100          105          110

Glu Thr Ile Leu Ser Leu Arg Phe Leu Ile Phe Gln Tyr Gly Ile Val
           115          120          125
    
```

-continued

Tyr Lys Leu Lys Ile Ala His
130 135

<210> SEQ ID NO 17
 <211> LENGTH: 306
 <212> TYPE: DNA
 <213> ORGANISM: Artificial sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Synthetic construct
 <220> FEATURE:
 <221> NAME/KEY: CDS
 <222> LOCATION: (1)..(306)
 <223> OTHER INFORMATION: Oryza sativa (rice) portion of 1,3-beta-D-glucan synthase (codon optimized)

<400> SEQUENCE: 17

tgg gtc gtt gct ttc gcc atc ctg tac aaa gaa gcc tgg aac aac cgc	48
Trp Val Val Ala Phe Ala Ile Leu Tyr Lys Glu Ala Trp Asn Asn Arg	
1 5 10 15	
aat tcg aat agc caa atc atg cgc ttt ttg tat gca gcc gcg gtg ttc	96
Asn Ser Asn Ser Gln Ile Met Arg Phe Leu Tyr Ala Ala Ala Val Phe	
20 25 30	
atg atc ccc gag gtc ctg gcg atc gtg ctg ttc atc gtc ccg tgg gtc	144
Met Ile Pro Glu Val Leu Ala Ile Val Leu Phe Ile Val Pro Trp Val	
35 40 45	
cgg aac gcc ctg gag aaa acc aat tgg aag att tgc tat gcg ctc acc	192
Arg Asn Ala Leu Glu Lys Thr Asn Trp Lys Ile Cys Tyr Ala Leu Thr	
50 55 60	
tgg tgg ttc cag agc cgc tcg ttc gtg ggt cgg ggc ctc cgc gag ggc	240
Trp Trp Phe Gln Ser Arg Ser Phe Val Gly Arg Gly Leu Arg Glu Gly	
65 70 75 80	
acg ttc gac aac gtg aag tac tcc gtg ttc tgg gtc ctt ctc ctc gcg	288
Thr Phe Asp Asn Val Lys Tyr Ser Val Phe Trp Val Leu Leu Leu Ala	
85 90 95	
gtc aag ttc gcg ttc tcc	306
Val Lys Phe Ala Phe Ser	
100	

<210> SEQ ID NO 18
 <211> LENGTH: 102
 <212> TYPE: PRT
 <213> ORGANISM: Artificial sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Synthetic Construct

<400> SEQUENCE: 18

Trp Val Val Ala Phe Ala Ile Leu Tyr Lys Glu Ala Trp Asn Asn Arg	
1 5 10 15	
Asn Ser Asn Ser Gln Ile Met Arg Phe Leu Tyr Ala Ala Ala Val Phe	
20 25 30	
Met Ile Pro Glu Val Leu Ala Ile Val Leu Phe Ile Val Pro Trp Val	
35 40 45	
Arg Asn Ala Leu Glu Lys Thr Asn Trp Lys Ile Cys Tyr Ala Leu Thr	
50 55 60	
Trp Trp Phe Gln Ser Arg Ser Phe Val Gly Arg Gly Leu Arg Glu Gly	
65 70 75 80	
Thr Phe Asp Asn Val Lys Tyr Ser Val Phe Trp Val Leu Leu Leu Ala	
85 90 95	
Val Lys Phe Ala Phe Ser	
100	

-continued

```

<210> SEQ ID NO 19
<211> LENGTH: 435
<212> TYPE: DNA
<213> ORGANISM: Artificial sequence
<220> FEATURE:
<223> OTHER INFORMATION: Synthetic construct
<220> FEATURE:
<221> NAME/KEY: CDS
<222> LOCATION: (1)..(435)
<223> OTHER INFORMATION: Glycine max (soy) portion of 1,3-beta-D-glucan
        synthase (codon optimized)

<400> SEQUENCE: 19

ttg agg tcc tcg gaa atg cgg aaa att ata gcc acg ctg cgc gct ctt      48
Leu Arg Ser Ser Glu Met Arg Lys Ile Ile Ala Thr Leu Arg Ala Leu
1           5           10           15

gtt gaa gtc ttg gaa tcc ctg tcg aag gac gcg gat ccg ggt ggc gtc      96
Val Glu Val Leu Glu Ser Leu Ser Lys Asp Ala Asp Pro Gly Gly Val
          20           25           30

ggt ggc ctg atc atg gaa gaa ctc cgg aag atc aag aaa tcg agt gtg     144
Gly Gly Leu Ile Met Glu Glu Leu Arg Lys Ile Lys Lys Ser Ser Val
          35           40           45

acc ctg tcg ggc gag ctc acc ccc tat aac atc att ccg ctt gag gcg     192
Thr Leu Ser Gly Glu Leu Thr Pro Tyr Asn Ile Ile Pro Leu Glu Ala
          50           55           60

ccg tcc ctc acc aat ccc atc cgg atc ttc ccc gag gtg aag gcc gcg     240
Pro Ser Leu Thr Asn Pro Ile Arg Ile Phe Pro Glu Val Lys Ala Ala
          65           70           75           80

atc agc gcg atc cgc tac acg gac cag ttc cca cgc ctc cct gcc ggc     288
Ile Ser Ala Ile Arg Tyr Thr Asp Gln Phe Pro Arg Leu Pro Ala Gly
          85           90           95

ttc aag atc agc ggg cag cgc gac gcg gat atg ttc gac ctc ctg gag     336
Phe Lys Ile Ser Gly Gln Arg Asp Ala Asp Met Phe Asp Leu Leu Glu
          100          105          110

ttc gtc ttt gga ttc cag aaa gac aac gtg cgc aac cag cgg gag aat     384
Phe Val Phe Gly Phe Gln Lys Asp Asn Val Arg Asn Gln Arg Glu Asn
          115          120          125

gtc gtg ctg atg atc gcc aac aag caa agc cgc ctc ggc atc ccg gca     432
Val Val Leu Met Ile Ala Asn Lys Gln Ser Arg Leu Gly Ile Pro Ala
          130          135          140

gag                                           435
Glu
145

```

```

<210> SEQ ID NO 20
<211> LENGTH: 145
<212> TYPE: PRT
<213> ORGANISM: Artificial sequence
<220> FEATURE:
<223> OTHER INFORMATION: Synthetic Construct

<400> SEQUENCE: 20

Leu Arg Ser Ser Glu Met Arg Lys Ile Ile Ala Thr Leu Arg Ala Leu
1           5           10           15

Val Glu Val Leu Glu Ser Leu Ser Lys Asp Ala Asp Pro Gly Gly Val
          20           25           30

Gly Gly Leu Ile Met Glu Glu Leu Arg Lys Ile Lys Lys Ser Ser Val
          35           40           45

Thr Leu Ser Gly Glu Leu Thr Pro Tyr Asn Ile Ile Pro Leu Glu Ala

```

-continued

50	55	60	
Pro Ser Leu Thr Asn Pro Ile Arg Ile Phe Pro Glu Val Lys Ala Ala			
65	70	75	80
Ile Ser Ala Ile Arg Tyr Thr Asp Gln Phe Pro Arg Leu Pro Ala Gly			
	85	90	95
Phe Lys Ile Ser Gly Gln Arg Asp Ala Asp Met Phe Asp Leu Leu Glu			
	100	105	110
Phe Val Phe Gly Phe Gln Lys Asp Asn Val Arg Asn Gln Arg Glu Asn			
	115	120	125
Val Val Leu Met Ile Ala Asn Lys Gln Ser Arg Leu Gly Ile Pro Ala			
	130	135	140
Glu			
145			
<210> SEQ ID NO 21			
<211> LENGTH: 489			
<212> TYPE: DNA			
<213> ORGANISM: Artificial sequence			
<220> FEATURE:			
<223> OTHER INFORMATION: Synthetic construct			
<220> FEATURE:			
<221> NAME/KEY: CDS			
<222> LOCATION: (1)..(489)			
<223> OTHER INFORMATION: Veronia mespilifolia 1,3-beta-D-glucan synthase (codon optimized)			
<400> SEQUENCE: 21			
tat ggc cat cca gat gtc ttt gac cgc gtt ttt cat att acc agg gga			48
Tyr Gly His Pro Asp Val Phe Asp Arg Val Phe His Ile Thr Arg Gly			
1	5	10	15
gga atc agt aaa gct agc cgc gtg atc aat atc tcc gaa gat atc tat			96
Gly Ile Ser Lys Ala Ser Arg Val Ile Asn Ile Ser Glu Asp Ile Tyr			
	20	25	30
gcc ggc ttc aat tcc acc ctg cgg caa ggc aat atc acc cac cac gag			144
Ala Gly Phe Asn Ser Thr Leu Arg Gln Gly Asn Ile Thr His His Glu			
	35	40	45
tat atc cag gtc ggt aag ggc cgc gac gtc ggc ctg aac cag att gcc			192
Tyr Ile Gln Val Gly Lys Gly Arg Asp Val Gly Leu Asn Gln Ile Ala			
	50	55	60
ctc ttc gag ggc aag gtc gcg ggc ggg aac ggc gag caa gtc ctc tcg			240
Leu Phe Glu Gly Lys Val Ala Gly Gly Asn Gly Glu Gln Val Leu Ser			
	65	70	75
cgc gac atc tac cgc ctc ggc cag ctg ttc gac ttc ttc cgg atg ctg			288
Arg Asp Ile Tyr Arg Leu Gly Gln Leu Phe Asp Phe Phe Arg Met Leu			
	85	90	95
tcg ttc tac ttc acg acc gtg ggg tac tat ttc tgc acc atg ctg acc			336
Ser Phe Tyr Phe Thr Thr Val Gly Tyr Tyr Phe Cys Thr Met Leu Thr			
	100	105	110
gtg acg act gtg tac ata ttc ctc tat ggt aag acc tac ttg gcc ctg			384
Val Thr Thr Val Tyr Ile Phe Leu Tyr Gly Lys Thr Tyr Leu Ala Leu			
	115	120	125
tcg ggt gtc ggc gag gac atc cag aac cgg agc gaa gtc ctc gac aac			432
Ser Gly Val Gly Glu Asp Ile Gln Asn Arg Ser Glu Val Leu Asp Asn			
	130	135	140
aaa gcg ctt acc gca gcg ctg aac acg cag ttc ctc ttc cag atc ggc			480
Lys Ala Leu Thr Ala Ala Leu Asn Thr Gln Phe Leu Phe Gln Ile Gly			
	145	150	155
gtg ttc acg			489

-continued

Val Phe Thr

<210> SEQ ID NO 22
 <211> LENGTH: 163
 <212> TYPE: PRT
 <213> ORGANISM: Artificial sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Synthetic Construct

<400> SEQUENCE: 22

```

Tyr Gly His Pro Asp Val Phe Asp Arg Val Phe His Ile Thr Arg Gly
1           5           10           15
Gly Ile Ser Lys Ala Ser Arg Val Ile Asn Ile Ser Glu Asp Ile Tyr
20           25           30
Ala Gly Phe Asn Ser Thr Leu Arg Gln Gly Asn Ile Thr His His Glu
35           40           45
Tyr Ile Gln Val Gly Lys Gly Arg Asp Val Gly Leu Asn Gln Ile Ala
50           55           60
Leu Phe Glu Gly Lys Val Ala Gly Gly Asn Gly Glu Gln Val Leu Ser
65           70           75           80
Arg Asp Ile Tyr Arg Leu Gly Gln Leu Phe Asp Phe Phe Arg Met Leu
85           90           95
Ser Phe Tyr Phe Thr Thr Val Gly Tyr Tyr Phe Cys Thr Met Leu Thr
100          105          110
Val Thr Thr Val Tyr Ile Phe Leu Tyr Gly Lys Thr Tyr Leu Ala Leu
115          120          125
Ser Gly Val Gly Glu Asp Ile Gln Asn Arg Ser Glu Val Leu Asp Asn
130          135          140
Lys Ala Leu Thr Ala Ala Leu Asn Thr Gln Phe Leu Phe Gln Ile Gly
145          150          155          160
    
```

Val Phe Thr

<210> SEQ ID NO 23
 <211> LENGTH: 1467
 <212> TYPE: DNA
 <213> ORGANISM: Artificial sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Synthetic construct
 <220> FEATURE:
 <221> NAME/KEY: CDS
 <222> LOCATION: (1)..(1467)
 <223> OTHER INFORMATION: Triticum aestivum (wheat) 1,3-beta-D-glucan
 synthase (codon optimized)

<400> SEQUENCE: 23

```

cgc gtg ggg aag ggt agg gat gtc ggc ttg aat caa atc agt atg ttc      48
Arg Val Gly Lys Gly Arg Asp Val Gly Leu Asn Gln Ile Ser Met Phe
1           5           10           15
gaa gca aaa gtg gct ggg gga aat ggt gag cag act ctg tcc cgc gac      96
Glu Ala Lys Val Ala Gly Gly Asn Gly Glu Gln Thr Leu Ser Arg Asp
20           25           30
gtc tac cgc ctt ggc cat ggc ctg gac ttc ttc cgg atg ctg agc ttc      144
Val Tyr Arg Leu Gly His Gly Leu Asp Phe Phe Arg Met Leu Ser Phe
35           40           45
ttc tac acg acc atc ggc ttt tac ctc aat acc atg atg gtc gtc ctc      192
Phe Tyr Thr Thr Ile Gly Phe Tyr Leu Asn Thr Met Met Val Val Leu
50           55           60
acg gtg tac gcc ttc gtg tgg ggt cgg ttc tac ctc gcg ctg tgg ggc      240
Thr Val Tyr Ala Phe Val Trp Gly Arg Phe Tyr Leu Ala Leu Ser Gly
    
```

-continued

65	70	75	80	
ctt gag gcc gac tat atc acc aat aac acc tcc tcg acc gat aat gcc				288
Leu Glu Ala Asp Tyr Ile Thr Asn Asn Thr Ser Ser Thr Asp Asn Ala	85	90	95	
gcg ctg tgg gca gtg ctc aac caa cag ttc ttc atc caa ttc ggc ctc				336
Ala Leu Trp Ala Val Leu Asn Gln Gln Phe Phe Ile Gln Phe Gly Leu	100	105	110	
ttc acg gcc ctc ccc atg atc atc gag aac tcc ctt gag cat ggc ttc				384
Phe Thr Ala Leu Pro Met Ile Ile Glu Asn Ser Leu Glu His Gly Phe	115	120	125	
ctc ata gcc gtc tgg gac ttc atc gtc atg cag ctg cag tgc gcg tcg				432
Leu Ile Ala Val Trp Asp Phe Ile Val Met Gln Leu Gln Cys Ala Ser	130	135	140	
gtg ttc tat acc ttc tgc atg ggc acc aag act cac tat tat ggc cgc				480
Val Phe Tyr Thr Phe Cys Met Gly Thr Lys Thr His Tyr Tyr Gly Arg	145	150	155	160
acg ctg ctg cat ggt ggg gcc aag tac cgc cca acc ggt cgg ggc ttc				528
Thr Leu Leu His Gly Gly Ala Lys Tyr Arg Pro Thr Gly Arg Gly Phe	165	170	175	
gtg gtc gag cac aag aaa ttc gcc gag aac tac cgg ctg tat gcg cgc				576
Val Val Glu His Lys Lys Phe Ala Glu Asn Tyr Arg Leu Tyr Ala Arg	180	185	190	
agc cat ttc acc aag gct atc gag ctg ggc gtg atc ttg tgt ttg tat				624
Ser His Phe Thr Lys Ala Ile Glu Leu Gly Val Ile Leu Cys Leu Tyr	195	200	205	
tcc tcg tac agc aac atc gct ggc gac acc ctg gtg tat att ctg ctg				672
Ser Ser Tyr Ser Asn Ile Ala Gly Asp Thr Leu Val Tyr Ile Leu Leu	210	215	220	
acc ctc tcg tcg tgg ttt ctc gtc tgc tcc tgg atc ctc gcg ccg ttc				720
Thr Leu Ser Ser Trp Phe Leu Val Cys Ser Trp Ile Leu Ala Pro Phe	225	230	235	240
atc ttc aac ccg agc gga ctc gat tgg cag aag aat tcc aac gac ttc				768
Ile Phe Asn Pro Ser Gly Leu Asp Trp Gln Lys Asn Ser Asn Asp Phe	245	250	255	
gag gat ttc ttc tcg tgg atc tgg ttt cag ggc ggc ggc atc agt gtc				816
Glu Asp Phe Phe Ser Trp Ile Trp Phe Gln Gly Gly Gly Ile Ser Val	260	265	270	
aag tcc gac cag agc tgg gag aag tgg tgg gag gaa gaa acc gac cat				864
Lys Ser Asp Gln Ser Trp Glu Lys Trp Trp Glu Glu Glu Thr Asp His	275	280	285	
ctg gcg cgg acg acg acc ggc ctg tgg ggc agc atc atc gag ata att				912
Leu Ala Arg Thr Thr Thr Gly Leu Trp Gly Ser Ile Ile Glu Ile Ile	290	295	300	
ctg gac ctg gcg cgc acc tac ttt ttc ttc cag tat gcg atc gtt tat				960
Leu Asp Leu Ala Arg Thr Tyr Phe Phe Phe Gln Tyr Ala Ile Val Tyr	305	310	315	320
cgc ctc cac atg gcc ggt ggc agc cgc tcc atc ctg gtc tat gtg ctc				1008
Arg Leu His Met Ala Gly Gly Ser Arg Ser Ile Leu Val Tyr Val Leu	325	330	335	
tcg tgg gcg tgc atc ccg ctc ccg ttc ctg gcg ctg gtc acc gtg acg				1056
Ser Trp Ala Cys Ile Pro Leu Pro Phe Leu Ala Leu Val Thr Val Thr	340	345	350	
tac ttc cgc gac aag tac tcg gcc aag aaa cat atc cgg tac cgc ctt				1104
Tyr Phe Arg Asp Lys Tyr Ser Ala Lys Lys His Ile Arg Tyr Arg Leu	355	360	365	
gtt caa tcg gtg att gtc tgc gca agc ctg gcg gcg att atc gtg ctc				1152
Val Gln Ser Val Ile Val Cys Ala Ser Leu Ala Ala Ile Ile Val Leu				

-continued

370	375	380	
ctc acc ctc acc aag ttc cag ttc atc gac acc ttc acc agc ctc ctg			1200
Leu Thr Leu Thr Lys Phe Gln Phe Ile Asp Thr Phe Thr Ser Leu Leu			
385	390	395	400
gcc ttt ctg ccc acc ggc tgg ggc atc atc tcc atc gcc ctg gtg ttc			1248
Ala Phe Leu Pro Thr Gly Trp Gly Ile Ile Ser Ile Ala Leu Val Phe			
	405	410	415
cgc caa tat ctg aag aaa agc gac acg gtg tgg aaa acc gtc gtc gtg			1296
Arg Gln Tyr Leu Lys Lys Ser Asp Thr Val Trp Lys Thr Val Val Val			
	420	425	430
gtc gcg cgg ttc tac gac atc acc ctc ggc ctg att gtt atg gcg ccg			1344
Val Ala Arg Phe Tyr Asp Ile Thr Leu Gly Leu Ile Val Met Ala Pro			
	435	440	445
atc gtc gtc ctg tcg tgg ctc cct ggc ctc cgc gag ctg cag acg cgg			1392
Ile Val Val Leu Ser Trp Leu Pro Gly Leu Arg Glu Leu Gln Thr Arg			
	450	455	460
atc ttg ttc aac gaa gcc ttc tcc aag ggc ctc cac atc tcg cag atg			1440
Ile Leu Phe Asn Glu Ala Phe Ser Lys Gly Leu His Ile Ser Gln Met			
	465	470	475
atc acg cgc agg aaa acg cat cgc gcc			1467
Ile Thr Arg Arg Lys Thr His Arg Ala			
	485		
<210> SEQ ID NO 24			
<211> LENGTH: 489			
<212> TYPE: PRT			
<213> ORGANISM: Artificial sequence			
<220> FEATURE:			
<223> OTHER INFORMATION: Synthetic Construct			
<400> SEQUENCE: 24			
Arg Val Gly Lys Gly Arg Asp Val Gly Leu Asn Gln Ile Ser Met Phe			
1	5	10	15
Glu Ala Lys Val Ala Gly Gly Asn Gly Glu Gln Thr Leu Ser Arg Asp			
	20	25	30
Val Tyr Arg Leu Gly His Gly Leu Asp Phe Phe Arg Met Leu Ser Phe			
	35	40	45
Phe Tyr Thr Thr Ile Gly Phe Tyr Leu Asn Thr Met Met Val Val Leu			
	50	55	60
Thr Val Tyr Ala Phe Val Trp Gly Arg Phe Tyr Leu Ala Leu Ser Gly			
	65	70	75
Leu Glu Ala Asp Tyr Ile Thr Asn Asn Thr Ser Ser Thr Asp Asn Ala			
	85	90	95
Ala Leu Trp Ala Val Leu Asn Gln Gln Phe Phe Ile Gln Phe Gly Leu			
	100	105	110
Phe Thr Ala Leu Pro Met Ile Ile Glu Asn Ser Leu Glu His Gly Phe			
	115	120	125
Leu Ile Ala Val Trp Asp Phe Ile Val Met Gln Leu Gln Cys Ala Ser			
	130	135	140
Val Phe Tyr Thr Phe Cys Met Gly Thr Lys Thr His Tyr Tyr Gly Arg			
	145	150	155
Thr Leu Leu His Gly Gly Ala Lys Tyr Arg Pro Thr Gly Arg Gly Phe			
	165	170	175
Val Val Glu His Lys Lys Phe Ala Glu Asn Tyr Arg Leu Tyr Ala Arg			
	180	185	190

-continued

Ser His Phe Thr Lys Ala Ile Glu Leu Gly Val Ile Leu Cys Leu Tyr
 195 200 205

Ser Ser Tyr Ser Asn Ile Ala Gly Asp Thr Leu Val Tyr Ile Leu Leu
 210 215 220

Thr Leu Ser Ser Trp Phe Leu Val Cys Ser Trp Ile Leu Ala Pro Phe
 225 230 235 240

Ile Phe Asn Pro Ser Gly Leu Asp Trp Gln Lys Asn Ser Asn Asp Phe
 245 250 255

Glu Asp Phe Phe Ser Trp Ile Trp Phe Gln Gly Gly Gly Ile Ser Val
 260 265 270

Lys Ser Asp Gln Ser Trp Glu Lys Trp Trp Glu Glu Glu Thr Asp His
 275 280 285

Leu Ala Arg Thr Thr Thr Gly Leu Trp Gly Ser Ile Ile Glu Ile Ile
 290 295 300

Leu Asp Leu Ala Arg Thr Tyr Phe Phe Phe Gln Tyr Ala Ile Val Tyr
 305 310 315 320

Arg Leu His Met Ala Gly Gly Ser Arg Ser Ile Leu Val Tyr Val Leu
 325 330 335

Ser Trp Ala Cys Ile Pro Leu Pro Phe Leu Ala Leu Val Thr Val Thr
 340 345 350

Tyr Phe Arg Asp Lys Tyr Ser Ala Lys Lys His Ile Arg Tyr Arg Leu
 355 360 365

Val Gln Ser Val Ile Val Cys Ala Ser Leu Ala Ala Ile Ile Val Leu
 370 375 380

Leu Thr Leu Thr Lys Phe Gln Phe Ile Asp Thr Phe Thr Ser Leu Leu
 385 390 395 400

Ala Phe Leu Pro Thr Gly Trp Gly Ile Ile Ser Ile Ala Leu Val Phe
 405 410 415

Arg Gln Tyr Leu Lys Lys Ser Asp Thr Val Trp Lys Thr Val Val Val
 420 425 430

Val Ala Arg Phe Tyr Asp Ile Thr Leu Gly Leu Ile Val Met Ala Pro
 435 440 445

Ile Val Val Leu Ser Trp Leu Pro Gly Leu Arg Glu Leu Gln Thr Arg
 450 455 460

Ile Leu Phe Asn Glu Ala Phe Ser Lys Gly Leu His Ile Ser Gln Met
 465 470 475 480

Ile Thr Arg Arg Lys Thr His Arg Ala
 485

<210> SEQ ID NO 25
 <211> LENGTH: 336
 <212> TYPE: DNA
 <213> ORGANISM: Artificial sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Synthetic construct
 <220> FEATURE:
 <221> NAME/KEY: CDS
 <222> LOCATION: (1)..(336)
 <223> OTHER INFORMATION: Hordeum vulgare (barley) 1,3-beta-D-glucan
 synthase (codon optimized)

<400> SEQUENCE: 25

att gcc gga gca gcc gct gga atc gct gga acc ctc atg tgt cac ccc 48
 Ile Ala Gly Ala Ala Ala Gly Ile Ala Gly Thr Leu Met Cys His Pro
 1 5 10 15

-continued

```

ctc gaa gtc ata aaa gat cgc ttg acc gtc gac agg gtg acg tat ccg      96
Leu Glu Val Ile Lys Asp Arg Leu Thr Val Asp Arg Val Thr Tyr Pro
      20                25                30

tcg atc agc atc gcg ttc tcg aag atc tac cgg acc gag ggc atc cgc      144
Ser Ile Ser Ile Ala Phe Ser Lys Ile Tyr Arg Thr Glu Gly Ile Arg
      35                40                45

ggc ctg tat agc ggc ctg tgc ccc acc ctc att ggc atg ctg ccg tac      192
Gly Leu Tyr Ser Gly Leu Cys Pro Thr Leu Ile Gly Met Leu Pro Tyr
      50                55                60

tcg act tgc tac tat ttc atg tat gac acg atc aag acc tcc tac tgc      240
Ser Thr Cys Tyr Tyr Phe Met Tyr Asp Thr Ile Lys Thr Ser Tyr Cys
      65                70                75                80

cgg ctc cat aag aaa aag tcg ctg agt cgc cct gag ctg ctg atc atc      288
Arg Leu His Lys Lys Lys Ser Leu Ser Arg Pro Glu Leu Leu Ile Ile
      85                90                95

ggt gcg ctt acc agc ctc acc gcg tcc acg atc tcc ttc ccg ctg gag      336
Gly Ala Leu Thr Ser Leu Thr Ala Ser Thr Ile Ser Phe Pro Leu Glu
      100                105                110

```

```

<210> SEQ ID NO 26
<211> LENGTH: 112
<212> TYPE: PRT
<213> ORGANISM: Artificial sequence
<220> FEATURE:
<223> OTHER INFORMATION: Synthetic Construct

```

```

<400> SEQUENCE: 26

```

```

Ile Ala Gly Ala Ala Ala Gly Ile Ala Gly Thr Leu Met Cys His Pro
1          5          10          15

Leu Glu Val Ile Lys Asp Arg Leu Thr Val Asp Arg Val Thr Tyr Pro
      20                25                30

Ser Ile Ser Ile Ala Phe Ser Lys Ile Tyr Arg Thr Glu Gly Ile Arg
      35                40                45

Gly Leu Tyr Ser Gly Leu Cys Pro Thr Leu Ile Gly Met Leu Pro Tyr
      50                55                60

Ser Thr Cys Tyr Tyr Phe Met Tyr Asp Thr Ile Lys Thr Ser Tyr Cys
      65                70                75                80

Arg Leu His Lys Lys Lys Ser Leu Ser Arg Pro Glu Leu Leu Ile Ile
      85                90                95

Gly Ala Leu Thr Ser Leu Thr Ala Ser Thr Ile Ser Phe Pro Leu Glu
      100                105                110

```

```

<210> SEQ ID NO 27
<211> LENGTH: 1293
<212> TYPE: DNA
<213> ORGANISM: Artificial sequence
<220> FEATURE:
<223> OTHER INFORMATION: Synthetic construct
<220> FEATURE:
<221> NAME/KEY: CDS
<222> LOCATION: (1)..(1293)
<223> OTHER INFORMATION: E.coli Glucose-1-phosphate adenylyltransferase
      (Acc. No. YP 49003.1) (codon optimized)

```

```

<400> SEQUENCE: 27

```

```

atg gtt tcc ctg gag aaa aat gac cac ctg atg ctc gca cgc caa ctc      48
Met Val Ser Leu Glu Lys Asn Asp His Leu Met Leu Ala Arg Gln Leu
1          5          10          15

ccg ctt aag tcc gtc gcc ctg atc ctc gcc ggc gga cgc ggc acg cgg      96
Pro Leu Lys Ser Val Ala Leu Ile Leu Ala Gly Gly Arg Gly Thr Arg

```

-continued

20		25		30												
ctc	aaa	gac	ctc	acc	aac	aag	cgc	gcg	aaa	ccg	gct	gtc	cat	ttc	ggt	144
Leu	Lys	Asp	Leu	Thr	Asn	Lys	Arg	Ala	Lys	Pro	Ala	Val	His	Phe	Gly	
	35						40					45				
ggc	aag	ttc	agg	atc	ata	gac	ttc	gcg	ctg	tcg	aac	tgc	atc	aat	tcc	192
Gly	Lys	Phe	Arg	Ile	Ile	Asp	Phe	Ala	Leu	Ser	Asn	Cys	Ile	Asn	Ser	
	50					55					60					
ggc	att	agg	cgc	atg	gga	gtc	att	acc	cag	tac	caa	tcg	cat	acg	ctc	240
Gly	Ile	Arg	Arg	Met	Gly	Val	Ile	Thr	Gln	Tyr	Gln	Ser	His	Thr	Leu	
	65				70					75					80	
gtc	cag	cat	atc	cag	cgg	ggc	tgg	tcg	ttc	ttc	aac	gaa	gag	atg	aac	288
Val	Gln	His	Ile	Gln	Arg	Gly	Trp	Ser	Phe	Phe	Asn	Glu	Glu	Met	Asn	
			85						90						95	
gag	ttc	gtc	gac	ctc	ctc	ccg	gcg	cag	cag	cgg	atg	aaa	ggc	gag	aac	336
Glu	Phe	Val	Asp	Leu	Leu	Pro	Ala	Gln	Gln	Arg	Met	Lys	Gly	Glu	Asn	
			100					105						110		
tgg	tac	cgc	ggc	acg	gct	gat	gcc	ggt	acc	cag	aac	ctg	gac	att	att	384
Trp	Tyr	Arg	Gly	Thr	Ala	Asp	Ala	Val	Thr	Gln	Asn	Leu	Asp	Ile	Ile	
			115					120						125		
cgc	cgc	tat	aaa	gcc	gag	tat	ggt	gtg	atc	ctg	gcc	ggt	gac	cac	atc	432
Arg	Arg	Tyr	Lys	Ala	Glu	Tyr	Val	Val	Ile	Leu	Ala	Gly	Asp	His	Ile	
		130					135						140			
tac	aaa	caa	gac	tat	agt	cgg	atg	ctc	atc	gac	cat	gtg	gaa	aag	ggc	480
Tyr	Lys	Gln	Asp	Tyr	Ser	Arg	Met	Leu	Ile	Asp	His	Val	Glu	Lys	Gly	
					150					155					160	
gct	cgc	tgc	acc	gtg	gcg	tgc	atg	cca	gtg	ccg	atc	gaa	gag	gcc	tcc	528
Ala	Arg	Cys	Thr	Val	Ala	Cys	Met	Pro	Val	Pro	Ile	Glu	Glu	Ala	Ser	
				165					170						175	
gcg	ttc	ggc	gtg	atg	gcc	gtg	gat	gag	aac	gac	aag	atc	atc	gag	ttc	576
Ala	Phe	Gly	Val	Met	Ala	Val	Asp	Glu	Asn	Asp	Lys	Ile	Ile	Glu	Phe	
			180						185					190		
gtg	gag	aag	ccc	gcg	aac	ccg	ccg	tcg	atg	ccc	aac	gac	ccg	agc	aag	624
Val	Glu	Lys	Pro	Ala	Asn	Pro	Pro	Ser	Met	Pro	Asn	Asp	Pro	Ser	Lys	
			195					200						205		
agc	ctg	gcg	tcc	atg	ggc	atc	tac	gtc	ttt	gac	gcg	gat	tat	ctg	tac	672
Ser	Leu	Ala	Ser	Met	Gly	Ile	Tyr	Val	Phe	Asp	Ala	Asp	Tyr	Leu	Tyr	
			210					215						220		
gag	ctt	ttg	gaa	gag	gat	gat	cgg	gac	gag	aat	agc	tcg	cac	gac	ttc	720
Glu	Leu	Leu	Glu	Glu	Asp	Asp	Arg	Asp	Glu	Asn	Ser	Ser	His	Asp	Phe	
					230					235					240	
ggc	aaa	gac	ctg	atc	ccg	aag	atc	acc	gaa	gcc	ggg	ctg	gcg	tat	gcc	768
Gly	Lys	Asp	Leu	Ile	Pro	Lys	Ile	Thr	Glu	Ala	Gly	Leu	Ala	Tyr	Ala	
				245						250					255	
cat	cct	ttt	ccg	ctc	agc	tgc	gtg	cag	tcg	gac	ccc	gat	gcg	gag	ccg	816
His	Pro	Phe	Pro	Leu	Ser	Cys	Val	Gln	Ser	Asp	Pro	Asp	Ala	Glu	Pro	
			260						265						270	
tat	tgg	cgc	gac	gtg	ggt	acc	ctg	gaa	gcg	tac	tgg	aag	gcc	aat	ctc	864
Tyr	Trp	Arg	Asp	Val	Gly	Thr	Leu	Glu	Ala	Tyr	Trp	Lys	Ala	Asn	Leu	
			275					280						285		
gac	ctc	gcc	agc	gtg	gtg	ccg	gaa	ctg	gac	atg	tac	gac	cgc	aat	tgg	912
Asp	Leu	Ala	Ser	Val	Val	Pro	Glu	Leu	Asp	Met	Tyr	Asp	Arg	Asn	Trp	
			290					295							300	
ccg	atc	cgc	act	tac	aac	gag	agc	ctg	ccc	ccg	gcg	aag	ttc	gtc	cag	960
Pro	Ile	Arg	Thr	Tyr	Asn	Glu	Ser	Leu	Pro	Pro	Ala	Lys	Phe	Val	Gln	
					310						315				320	
gac	cgg	agt	ggc	agc	cac	ggc	atg	acg	ctc	aat	tcc	ctt	gtg	tcg	ggg	1008
Asp	Arg	Ser	Gly	Ser	His	Gly	Met	Thr	Leu	Asn	Ser	Leu	Val	Ser	Gly	

-continued

325	330	335	
ggc tgc gtc atc tcg ggt tcg gtc gtc gtc cag tcc gtc ctc ttc agc Gly Cys Val Ile Ser Gly Ser Val Val Val Gln Ser Val Leu Phe Ser 340 345 350			1056
cgg gtc agg gtc aat tcc ttc tgc aac atc gat agc gca gtg ctg ttg Arg Val Arg Val Asn Ser Phe Cys Asn Ile Asp Ser Ala Val Leu Leu 355 360 365			1104
ccc gag gtc tgg gtg ggc cgc tcg tgt cgg ctg cgc cgc tgc gtg atc Pro Glu Val Trp Val Gly Arg Ser Cys Arg Leu Arg Arg Cys Val Ile 370 375 380			1152
gac cgc gcc tgc gtc atc ccc gag ggc atg gtc ata ggc gag aat gcc Asp Arg Ala Cys Val Ile Pro Glu Gly Met Val Ile Gly Glu Asn Ala 385 390 395 400			1200
gaa gag gac gcg cgg cgc ttc tat cgg tcc gag gag ggc atc gtg ctg Glu Glu Asp Ala Arg Arg Phe Tyr Arg Ser Glu Glu Gly Ile Val Leu 405 410 415			1248
gtc acc cgc gag atg ctg cgc aag ctc ggg cat aag caa gag cgc Val Thr Arg Glu Met Leu Arg Lys Leu Gly His Lys Gln Glu Arg 420 425 430			1293

<210> SEQ ID NO 28
 <211> LENGTH: 431
 <212> TYPE: PRT
 <213> ORGANISM: Artificial sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Synthetic Construct

<400> SEQUENCE: 28

Met Val Ser Leu Glu Lys Asn Asp His Leu Met Leu Ala Arg Gln Leu 1 5 10 15
Pro Leu Lys Ser Val Ala Leu Ile Leu Ala Gly Gly Arg Gly Thr Arg 20 25 30
Leu Lys Asp Leu Thr Asn Lys Arg Ala Lys Pro Ala Val His Phe Gly 35 40 45
Gly Lys Phe Arg Ile Ile Asp Phe Ala Leu Ser Asn Cys Ile Asn Ser 50 55 60
Gly Ile Arg Arg Met Gly Val Ile Thr Gln Tyr Gln Ser His Thr Leu 65 70 75 80
Val Gln His Ile Gln Arg Gly Trp Ser Phe Phe Asn Glu Glu Met Asn 85 90 95
Glu Phe Val Asp Leu Leu Pro Ala Gln Gln Arg Met Lys Gly Glu Asn 100 105 110
Trp Tyr Arg Gly Thr Ala Asp Ala Val Thr Gln Asn Leu Asp Ile Ile 115 120 125
Arg Arg Tyr Lys Ala Glu Tyr Val Val Ile Leu Ala Gly Asp His Ile 130 135 140
Tyr Lys Gln Asp Tyr Ser Arg Met Leu Ile Asp His Val Glu Lys Gly 145 150 155 160
Ala Arg Cys Thr Val Ala Cys Met Pro Val Pro Ile Glu Glu Ala Ser 165 170 175
Ala Phe Gly Val Met Ala Val Asp Glu Asn Asp Lys Ile Ile Glu Phe 180 185 190
Val Glu Lys Pro Ala Asn Pro Pro Ser Met Pro Asn Asp Pro Ser Lys 195 200 205
Ser Leu Ala Ser Met Gly Ile Tyr Val Phe Asp Ala Asp Tyr Leu Tyr

-continued

210	215	220	
Glu Leu Leu Glu Glu Asp Asp Arg Asp Glu Asn Ser Ser His Asp Phe			
225	230	235	240
Gly Lys Asp Leu Ile Pro Lys Ile Thr Glu Ala Gly Leu Ala Tyr Ala			
	245	250	255
His Pro Phe Pro Leu Ser Cys Val Gln Ser Asp Pro Asp Ala Glu Pro			
	260	265	270
Tyr Trp Arg Asp Val Gly Thr Leu Glu Ala Tyr Trp Lys Ala Asn Leu			
	275	280	285
Asp Leu Ala Ser Val Val Pro Glu Leu Asp Met Tyr Asp Arg Asn Trp			
	290	295	300
Pro Ile Arg Thr Tyr Asn Glu Ser Leu Pro Pro Ala Lys Phe Val Gln			
	305	310	315
Asp Arg Ser Gly Ser His Gly Met Thr Leu Asn Ser Leu Val Ser Gly			
	325	330	335
Gly Cys Val Ile Ser Gly Ser Val Val Val Gln Ser Val Leu Phe Ser			
	340	345	350
Arg Val Arg Val Asn Ser Phe Cys Asn Ile Asp Ser Ala Val Leu Leu			
	355	360	365
Pro Glu Val Trp Val Gly Arg Ser Cys Arg Leu Arg Arg Cys Val Ile			
	370	375	380
Asp Arg Ala Cys Val Ile Pro Glu Gly Met Val Ile Gly Glu Asn Ala			
	385	390	395
Glu Glu Asp Ala Arg Arg Phe Tyr Arg Ser Glu Glu Gly Ile Val Leu			
	405	410	415
Val Thr Arg Glu Met Leu Arg Lys Leu Gly His Lys Gln Glu Arg			
	420	425	430

<210> SEQ ID NO 29
 <211> LENGTH: 1227
 <212> TYPE: DNA
 <213> ORGANISM: Artificial sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Synthetic construct
 <220> FEATURE:
 <221> NAME/KEY: CDS
 <222> LOCATION: (1)..(1227)
 <223> OTHER INFORMATION: Cornebacterium glutamicum (ATCC 13032)
 Glucose-1-phosphate adenylyltransferase (codon optimized)

<400> SEQUENCE: 29

atg gtg aag gga gtt aag gga agg cct aat gtt ttg gca ata gtt ctg	48
Met Val Lys Gly Val Lys Gly Arg Pro Asn Val Leu Ala Ile Val Leu	
1 5 10 15	
gcg ggt gga gag ggg aaa cgg ttg ttc ccg ctc acc gag gac cgc gcc	96
Ala Gly Gly Glu Gly Lys Arg Leu Phe Pro Leu Thr Glu Asp Arg Ala	
20 25 30	
aag ccc gcg gtg ccg ttc ggc ggc acc tac cgc ctg atc gat ttc gtg	144
Lys Pro Ala Val Pro Phe Gly Gly Thr Tyr Arg Leu Ile Asp Phe Val	
35 40 45	
ctg tcc aat ctg gtc aat tcg ggt ttc ctc aag atc gcg gtc ctc acg	192
Leu Ser Asn Leu Val Asn Ser Gly Phe Leu Lys Ile Ala Val Leu Thr	
50 55 60	
cag tac aag agc cat agc ctt gac cgg cat atc tcc ctg tcc tgg aac	240
Gln Tyr Lys Ser His Ser Leu Asp Arg His Ile Ser Leu Ser Trp Asn	
65 70 75 80	

-continued

gtg tcc ggg ccg acg ggc cag tac atc gcc tcc gtc cca gct cag cag	288
Val Ser Gly Pro Thr Gly Gln Tyr Ile Ala Ser Val Pro Ala Gln Gln	
85 90 95	
cgg ctc ggc aag cgc tgg ttc acc ggc tcg gcc gac gcc atc ctg cag	336
Arg Leu Gly Lys Arg Trp Phe Thr Gly Ser Ala Asp Ala Ile Leu Gln	
100 105 110	
agc ctc aac ctg atc tcc gac gag aag ccc gac tat gtc atc gtg ttt	384
Ser Leu Asn Leu Ile Ser Asp Glu Lys Pro Asp Tyr Val Ile Val Phe	
115 120 125	
ggc gcg gac cac gtg tac cgg atg gat ccc tcc cag atg ctg gat gag	432
Gly Ala Asp His Val Tyr Arg Met Asp Pro Ser Gln Met Leu Asp Glu	
130 135 140	
cat atc gcg agt ggt cgc gct gtg tcg gtc gcc ggc atc cgc gtc ccg	480
His Ile Ala Ser Gly Arg Ala Val Ser Val Ala Gly Ile Arg Val Pro	
145 150 155 160	
cgc gaa gag gcg acg gcg ttc ggc tgc atc cag tcc gat gtg gac ggg	528
Arg Glu Glu Ala Thr Ala Phe Gly Cys Ile Gln Ser Asp Val Asp Gly	
165 170 175	
aac atc acc gag ttc ctc gaa aaa ccc gcc gac ccc ccg ggg acc ccg	576
Asn Ile Thr Glu Phe Leu Glu Lys Pro Ala Asp Pro Pro Gly Thr Pro	
180 185 190	
gac gac ccc gac atg acc tat gcc agc atg gcc aac tac atc ttc acg	624
Asp Asp Pro Asp Met Thr Tyr Ala Ser Met Gly Asn Tyr Ile Phe Thr	
195 200 205	
acc gaa gca ctg atc caa gcg ctt aaa gat gat gag aat aac gaa aat	672
Thr Glu Ala Leu Ile Gln Ala Leu Lys Asp Asp Glu Asn Asn Glu Asn	
210 215 220	
tcg gac cat gac atg ggc ggc gac atc att ccg tat ttc gtg tcg cgc	720
Ser Asp His Asp Met Gly Gly Asp Ile Ile Pro Tyr Phe Val Ser Arg	
225 230 235 240	
aac gac gcg cat gtc tac gac ttt tcc ggt aac atc gtg ccg ggt gcg	768
Asn Asp Ala His Val Tyr Asp Phe Ser Gly Asn Ile Val Pro Gly Ala	
245 250 255	
act gag cgc gac aag ggc tat tgg cgc gac gtc ggt acc att gat gcg	816
Thr Glu Arg Asp Lys Gly Tyr Trp Arg Asp Val Gly Thr Ile Asp Ala	
260 265 270	
ttc tac gag tgc cac atg gac ctg atc tcg gtc cac ccg atc ttc aat	864
Phe Tyr Glu Cys His Met Asp Leu Ile Ser Val His Pro Ile Phe Asn	
275 280 285	
ctg tat aac agc gag tgg ccg atc cac acc acg tcc gag ggc aac ctc	912
Leu Tyr Asn Ser Glu Trp Pro Ile His Thr Thr Ser Glu Gly Asn Leu	
290 295 300	
ccg ccg gcc aag ttc gtc cgc ggc ggc ata gcc caa tcg tcg atg gtg	960
Pro Pro Ala Lys Phe Val Arg Gly Gly Ile Ala Gln Ser Ser Met Val	
305 310 315 320	
agc tcc ggc agc atc atc tcg gct ggc acc gtg agg aat agc gtg ctc	1008
Ser Ser Gly Ser Ile Ile Ser Ala Gly Thr Val Arg Asn Ser Val Leu	
325 330 335	
tcg aat aat gtc gtc gtc gag gag ggc gcc acg gtc gag ggc gcg gtg	1056
Ser Asn Asn Val Val Val Glu Glu Gly Ala Thr Val Glu Gly Ala Val	
340 345 350	
ctc atg ccc ggt gtc cgg att ggc aag ggt gcc gtc gtg cgc cat gca	1104
Leu Met Pro Gly Val Arg Ile Gly Lys Gly Ala Val Val Arg His Ala	
355 360 365	
att ctc gac aaa aac gtc gtc gtg cgc gac ggc gag ctc atc ggc gtg	1152
Ile Leu Asp Lys Asn Val Val Val Arg Asp Gly Glu Leu Ile Gly Val	
370 375 380	

-continued

gat cag gtc cgg gac gcc cag cgc ttc aag gtc agt gcg ggc gga gtg 1200
 Asp Gln Val Arg Asp Ala Gln Arg Phe Lys Val Ser Ala Gly Gly Val
 385 390 395 400

gtc gtg gtc ggc aag aac caa gtc gtg 1227
 Val Val Val Gly Lys Asn Gln Val Val
 405

<210> SEQ ID NO 30
 <211> LENGTH: 409
 <212> TYPE: PRT
 <213> ORGANISM: Artificial sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Synthetic Construct

<400> SEQUENCE: 30

Met Val Lys Gly Val Lys Gly Arg Pro Asn Val Leu Ala Ile Val Leu
 1 5 10 15
 Ala Gly Gly Glu Gly Lys Arg Leu Phe Pro Leu Thr Glu Asp Arg Ala
 20 25 30
 Lys Pro Ala Val Pro Phe Gly Gly Thr Tyr Arg Leu Ile Asp Phe Val
 35 40 45
 Leu Ser Asn Leu Val Asn Ser Gly Phe Leu Lys Ile Ala Val Leu Thr
 50 55 60
 Gln Tyr Lys Ser His Ser Leu Asp Arg His Ile Ser Leu Ser Trp Asn
 65 70 75 80
 Val Ser Gly Pro Thr Gly Gln Tyr Ile Ala Ser Val Pro Ala Gln Gln
 85 90 95
 Arg Leu Gly Lys Arg Trp Phe Thr Gly Ser Ala Asp Ala Ile Leu Gln
 100 105 110
 Ser Leu Asn Leu Ile Ser Asp Glu Lys Pro Asp Tyr Val Ile Val Phe
 115 120 125
 Gly Ala Asp His Val Tyr Arg Met Asp Pro Ser Gln Met Leu Asp Glu
 130 135 140
 His Ile Ala Ser Gly Arg Ala Val Ser Val Ala Gly Ile Arg Val Pro
 145 150 155 160
 Arg Glu Glu Ala Thr Ala Phe Gly Cys Ile Gln Ser Asp Val Asp Gly
 165 170 175
 Asn Ile Thr Glu Phe Leu Glu Lys Pro Ala Asp Pro Pro Gly Thr Pro
 180 185 190
 Asp Asp Pro Asp Met Thr Tyr Ala Ser Met Gly Asn Tyr Ile Phe Thr
 195 200 205
 Thr Glu Ala Leu Ile Gln Ala Leu Lys Asp Asp Glu Asn Asn Glu Asn
 210 215 220
 Ser Asp His Asp Met Gly Gly Asp Ile Ile Pro Tyr Phe Val Ser Arg
 225 230 235 240
 Asn Asp Ala His Val Tyr Asp Phe Ser Gly Asn Ile Val Pro Gly Ala
 245 250 255
 Thr Glu Arg Asp Lys Gly Tyr Trp Arg Asp Val Gly Thr Ile Asp Ala
 260 265 270
 Phe Tyr Glu Cys His Met Asp Leu Ile Ser Val His Pro Ile Phe Asn
 275 280 285
 Leu Tyr Asn Ser Glu Trp Pro Ile His Thr Thr Ser Glu Gly Asn Leu
 290 295 300
 Pro Pro Ala Lys Phe Val Arg Gly Gly Ile Ala Gln Ser Ser Met Val

-continued

305	310	315	320
Ser Ser Gly Ser Ile Ile Ser Ala Gly Thr Val Arg Asn Ser Val Leu			
	325	330	335
Ser Asn Asn Val Val Val Glu Glu Gly Ala Thr Val Glu Gly Ala Val			
	340	345	350
Leu Met Pro Gly Val Arg Ile Gly Lys Gly Ala Val Val Arg His Ala			
	355	360	365
Ile Leu Asp Lys Asn Val Val Val Arg Asp Gly Glu Leu Ile Gly Val			
	370	375	380
Asp Gln Val Arg Asp Ala Gln Arg Phe Lys Val Ser Ala Gly Gly Val			
	385	390	395
Val Val Val Gly Lys Asn Gln Val Val			
	405		

<210> SEQ ID NO 31
 <211> LENGTH: 1431
 <212> TYPE: DNA
 <213> ORGANISM: Artificial sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Synthetic construct
 <220> FEATURE:
 <221> NAME/KEY: CDS
 <222> LOCATION: (1)..(1431)
 <223> OTHER INFORMATION: Escherichia coli str. K-12 substr. W3110
 Glycogen Synthase (codon optimized)

<400> SEQUENCE: 31

atg caa gtt ctt cat gtg tgt tcc gaa atg ttc ccc ctc ctc aaa acc	48
Met Gln Val Leu His Val Cys Ser Glu Met Phe Pro Leu Leu Lys Thr	
1 5 10 15	
ggg ggc ctg gct gat gtc ata ggt gcc ctg ccg gct gcg cag att gcg	96
Gly Gly Leu Ala Asp Val Ile Gly Ala Leu Pro Ala Ala Gln Ile Ala	
20 25 30	
gac ggc gtg gac gca cgg gtc ctg ctg ccg gcg ttc ccg gat atc cgc	144
Asp Gly Val Asp Ala Arg Val Leu Leu Pro Ala Phe Pro Asp Ile Arg	
35 40 45	
cgg ggc gtg acc gac gca caa gtc gtg tcg cgc agg gac acc ttt gcc	192
Arg Gly Val Thr Asp Ala Gln Val Val Ser Arg Arg Asp Thr Phe Ala	
50 55 60	
ggc cac atc acg ctc ttg ttc ggc cac tat aac ggc gtg ggc atc tac	240
Gly His Ile Thr Leu Leu Phe Gly His Tyr Asn Gly Val Gly Ile Tyr	
65 70 75 80	
ctg atc gat gcg ccg cat ctc tat gac agg ccg ggt tcg ccc tat cat	288
Leu Ile Asp Ala Pro His Leu Tyr Asp Arg Pro Gly Ser Pro Tyr His	
85 90 95	
gac acg aac ctc ttc gcc tac acg gac aat gtg ctg ccg ttc gca ctt	336
Asp Thr Asn Leu Phe Ala Tyr Thr Asp Asn Val Leu Arg Phe Ala Leu	
100 105 110	
ctg ggc tgg gtc gga gcc gag atg gca tcc ggc ctc gac ccg ttc tgg	384
Leu Gly Trp Val Gly Ala Glu Met Ala Ser Gly Leu Asp Pro Phe Trp	
115 120 125	
cgc cct gac gtc gtc cat gcg cat gac tgg cat gcc ggc ctc gca ccc	432
Arg Pro Asp Val Val His Ala His Asp Trp His Ala Gly Leu Ala Pro	
130 135 140	
gcg tat ttg gcc gcc cgg gga cgg ccg gct aag agc gtg ttt acc gtt	480
Ala Tyr Leu Ala Ala Arg Gly Arg Pro Ala Lys Ser Val Phe Thr Val	
145 150 155 160	
cat aac ctc gcg tat cag ggc atg ttc tac gcc cat cac atg aat gat	528

-continued

His	Asn	Leu	Ala	Tyr	Gln	Gly	Met	Phe	Tyr	Ala	His	His	Met	Asn	Asp	
				165					170					175		
atc	cag	ctg	ccc	tgg	tcc	ttc	ttc	aac	atc	cac	ggg	ctt	gag	ttc	aat	576
Ile	Gln	Leu	Pro	Trp	Ser	Phe	Phe	Asn	Ile	His	Gly	Leu	Glu	Phe	Asn	
			180					185					190			
ggc	caa	atc	tcg	ttc	ctg	aag	gcc	ggg	ctg	tac	tac	gcg	gac	cac	atc	624
Gly	Gln	Ile	Ser	Phe	Leu	Lys	Ala	Gly	Leu	Tyr	Tyr	Ala	Asp	His	Ile	
			195				200					205				
acc	gcg	gtg	tcg	cca	acc	tac	gcc	cgc	gag	atc	acc	gag	ccg	cag	ttc	672
Thr	Ala	Val	Ser	Pro	Thr	Tyr	Ala	Arg	Glu	Ile	Thr	Glu	Pro	Gln	Phe	
			210			215					220					
gcg	tac	ggc	atg	gag	ggc	ctg	ctg	caa	cag	cgc	cac	cgc	gag	ggc	cgg	720
Ala	Tyr	Gly	Met	Glu	Gly	Leu	Leu	Gln	Gln	Arg	His	Arg	Glu	Gly	Arg	
					230					235					240	
ctc	agc	ggc	ggt	ctg	aac	ggc	gtc	gac	gag	aaa	atc	tgg	tcg	ccc	gag	768
Leu	Ser	Gly	Val	Leu	Asn	Gly	Val	Asp	Glu	Lys	Ile	Trp	Ser	Pro	Glu	
				245				250						255		
act	gac	ttg	ctc	ctt	gcc	agc	cgc	tat	acc	cgc	gac	acg	ctc	gaa	gat	816
Thr	Asp	Leu	Leu	Leu	Ala	Ser	Arg	Tyr	Thr	Arg	Asp	Thr	Leu	Glu	Asp	
				260				265					270			
aaa	gcc	gag	aat	aag	cgc	cag	ctg	cag	atc	gcc	atg	ggc	ctg	aaa	gtc	864
Lys	Ala	Glu	Asn	Lys	Arg	Gln	Leu	Gln	Ile	Ala	Met	Gly	Leu	Lys	Val	
				275			280					285				
gac	gac	aag	gtc	ccc	ctc	ttc	gcc	gtg	gtc	agc	cgc	ctg	acc	tcg	caa	912
Asp	Asp	Lys	Val	Pro	Leu	Phe	Ala	Val	Val	Ser	Arg	Leu	Thr	Ser	Gln	
				290		295				300						
aag	ggc	ctg	gac	ctg	gtg	ctc	gaa	gcc	ctc	cct	ggc	ctg	ctt	gaa	cag	960
Lys	Gly	Leu	Asp	Leu	Val	Leu	Glu	Ala	Leu	Pro	Gly	Leu	Leu	Glu	Gln	
					310					315					320	
ggg	ggc	cag	ttg	gcg	ctc	ctc	ggc	gcc	ggg	gat	ccg	gtg	ttg	cag	gag	1008
Gly	Gly	Gln	Leu	Ala	Leu	Leu	Gly	Ala	Gly	Asp	Pro	Val	Leu	Gln	Glu	
				325					330					335		
gga	ttc	ctg	gcg	gct	gcg	gcc	gag	tat	ccg	ggc	cag	gtc	ggc	gtc	cag	1056
Gly	Phe	Leu	Ala	Ala	Ala	Ala	Glu	Tyr	Pro	Gly	Gln	Val	Gly	Val	Gln	
				340				345						350		
att	ggc	tac	cat	gaa	gcg	ttc	agt	cat	cgg	atc	atg	ggg	ggg	gcc	gac	1104
Ile	Gly	Tyr	His	Glu	Ala	Phe	Ser	His	Arg	Ile	Met	Gly	Gly	Ala	Asp	
				355				360					365			
gtc	atc	ctc	gtg	ccg	tcc	cgc	ttc	gag	ccg	tgc	ggc	ctc	acc	cag	ctg	1152
Val	Ile	Leu	Val	Pro	Ser	Arg	Phe	Glu	Pro	Cys	Gly	Leu	Thr	Gln	Leu	
				370			375					380				
tac	ggc	ctc	aag	tac	gga	acg	ctc	ccc	ctg	gtg	cgg	cgg	acc	ggg	ggg	1200
Tyr	Gly	Leu	Lys	Tyr	Gly	Thr	Leu	Pro	Leu	Val	Arg	Arg	Thr	Gly	Gly	
				385		390				395					400	
ctc	gcc	gac	acc	gtc	agc	gac	tgc	tcc	ctg	gag	aac	ctg	gcg	gat	ggc	1248
Leu	Ala	Asp	Thr	Val	Ser	Asp	Cys	Ser	Leu	Glu	Asn	Leu	Ala	Asp	Gly	
				405					410					415		
gtc	gcg	agc	ggg	ttt	gtg	ttc	gag	gac	agc	aac	gcc	tgg	tcg	ctg	ctc	1296
Val	Ala	Ser	Gly	Phe	Val	Phe	Glu	Asp	Ser	Asn	Ala	Trp	Ser	Leu	Leu	
				420				425						430		
cgc	gcg	atc	cgc	agg	gcc	ttc	gtc	ctg	tgg	agt	cgc	ccg	tcc	ctc	tgg	1344
Arg	Ala	Ile	Arg	Arg	Ala	Phe	Val	Leu	Trp	Ser	Arg	Pro	Ser	Leu	Trp	
				435				440						445		
cgc	ttc	gtg	cag	cgg	cag	gca	atg	gcc	atg	gac	ttc	tcg	tgg	caa	gtc	1392
Arg	Phe	Val	Gln	Arg	Gln	Ala	Met	Ala	Met	Asp	Phe	Ser	Trp	Gln	Val	
				450			455				460					
gcg	gcc	aag	tcc	tat	cgc	gag	ctc	tac	tat	cgc	ctg	aag				1431

-continued

Ala Ala Lys Ser Tyr Arg Glu Leu Tyr Tyr Arg Leu Lys
465 470 475

<210> SEQ ID NO 32
 <211> LENGTH: 477
 <212> TYPE: PRT
 <213> ORGANISM: Artificial sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Synthetic Construct

<400> SEQUENCE: 32

Met Gln Val Leu His Val Cys Ser Glu Met Phe Pro Leu Leu Lys Thr
1 5 10 15
 Gly Gly Leu Ala Asp Val Ile Gly Ala Leu Pro Ala Ala Gln Ile Ala
20 25 30
 Asp Gly Val Asp Ala Arg Val Leu Leu Pro Ala Phe Pro Asp Ile Arg
35 40 45
 Arg Gly Val Thr Asp Ala Gln Val Val Ser Arg Arg Asp Thr Phe Ala
50 55 60
 Gly His Ile Thr Leu Leu Phe Gly His Tyr Asn Gly Val Gly Ile Tyr
65 70 75 80
 Leu Ile Asp Ala Pro His Leu Tyr Asp Arg Pro Gly Ser Pro Tyr His
85 90 95
 Asp Thr Asn Leu Phe Ala Tyr Thr Asp Asn Val Leu Arg Phe Ala Leu
100 105 110
 Leu Gly Trp Val Gly Ala Glu Met Ala Ser Gly Leu Asp Pro Phe Trp
115 120 125
 Arg Pro Asp Val Val His Ala His Asp Trp His Ala Gly Leu Ala Pro
130 135 140
 Ala Tyr Leu Ala Ala Arg Gly Arg Pro Ala Lys Ser Val Phe Thr Val
145 150 155 160
 His Asn Leu Ala Tyr Gln Gly Met Phe Tyr Ala His His Met Asn Asp
165 170 175
 Ile Gln Leu Pro Trp Ser Phe Phe Asn Ile His Gly Leu Glu Phe Asn
180 185 190
 Gly Gln Ile Ser Phe Leu Lys Ala Gly Leu Tyr Tyr Ala Asp His Ile
195 200 205
 Thr Ala Val Ser Pro Thr Tyr Ala Arg Glu Ile Thr Glu Pro Gln Phe
210 215 220
 Ala Tyr Gly Met Glu Gly Leu Leu Gln Gln Arg His Arg Glu Gly Arg
225 230 235 240
 Leu Ser Gly Val Leu Asn Gly Val Asp Glu Lys Ile Trp Ser Pro Glu
245 250 255
 Thr Asp Leu Leu Leu Ala Ser Arg Tyr Thr Arg Asp Thr Leu Glu Asp
260 265 270
 Lys Ala Glu Asn Lys Arg Gln Leu Gln Ile Ala Met Gly Leu Lys Val
275 280 285
 Asp Asp Lys Val Pro Leu Phe Ala Val Val Ser Arg Leu Thr Ser Gln
290 295 300
 Lys Gly Leu Asp Leu Val Leu Glu Ala Leu Pro Gly Leu Leu Glu Gln
305 310 315 320
 Gly Gly Gln Leu Ala Leu Leu Gly Ala Gly Asp Pro Val Leu Gln Glu
325 330 335

-continued

Gly Phe Leu Ala Ala Ala Ala Glu Tyr Pro Gly Gln Val Gly Val Gln
 340 345 350

Ile Gly Tyr His Glu Ala Phe Ser His Arg Ile Met Gly Gly Ala Asp
 355 360 365

Val Ile Leu Val Pro Ser Arg Phe Glu Pro Cys Gly Leu Thr Gln Leu
 370 375 380

Tyr Gly Leu Lys Tyr Gly Thr Leu Pro Leu Val Arg Arg Thr Gly Gly
 385 390 395 400

Leu Ala Asp Thr Val Ser Asp Cys Ser Leu Glu Asn Leu Ala Asp Gly
 405 410 415

Val Ala Ser Gly Phe Val Phe Glu Asp Ser Asn Ala Trp Ser Leu Leu
 420 425 430

Arg Ala Ile Arg Arg Ala Phe Val Leu Trp Ser Arg Pro Ser Leu Trp
 435 440 445

Arg Phe Val Gln Arg Gln Ala Met Ala Met Asp Phe Ser Trp Gln Val
 450 455 460

Ala Ala Lys Ser Tyr Arg Glu Leu Tyr Tyr Arg Leu Lys
 465 470 475

<210> SEQ ID NO 33
 <211> LENGTH: 1227
 <212> TYPE: DNA
 <213> ORGANISM: Artificial sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Synthetic construct
 <220> FEATURE:
 <221> NAME/KEY: CDS
 <222> LOCATION: (1)..(1227)
 <223> OTHER INFORMATION: Cornebacterium glutamicum (ATCC 13032)
 Glycosyltransferase (codon optimized)

<400> SEQUENCE: 33

atg cca cct ttc cgc tat cgc tgt gct act gtt ttc cgc tgg ttg att	48
Met Pro Pro Phe Arg Tyr Arg Cys Ala Thr Val Phe Arg Trp Leu Ile	
1 5 10 15	
ttt gaa ata atg cgc gtc ggg atg atg acc cgc gag tac ccg cct gaa	96
Phe Glu Ile Met Arg Val Gly Met Met Thr Arg Glu Tyr Pro Pro Glu	
20 25 30	
gtg tac ggt gga gcc ggg gtc cat gtg acc gag ctt acc cgg ttc atg	144
Val Tyr Gly Gly Ala Gly Val His Val Thr Glu Leu Thr Arg Phe Met	
35 40 45	
cgc gag atc gca gag gtc gac gtg cac tgc atg ggc gca ccg cgc gac	192
Arg Glu Ile Ala Glu Val Asp Val His Cys Met Gly Ala Pro Arg Asp	
50 55 60	
atg gag ggt gtt ttc gtg cac ggc gtc gac ccg gca ctc gaa tcg gcg	240
Met Glu Gly Val Phe Val His Gly Val Asp Pro Ala Leu Glu Ser Ala	
65 70 75 80	
aac ccc gcc atc aag acg ctc tcg acg ggc ctg cgg atg gcc gag gcc	288
Asn Pro Ala Ile Lys Thr Leu Ser Thr Gly Leu Arg Met Ala Glu Ala	
85 90 95	
gcg aat aat gtt gac gtc gtg cat tcg cat acc tgg tat gcg ggc ctg	336
Ala Asn Asn Val Asp Val Val His Ser His Thr Trp Tyr Ala Gly Leu	
100 105 110	
ggg ggc cat ctc gcg gcc agg ctg cac ggc atc ccg cat gtg gcg acg	384
Gly Gly His Leu Ala Ala Arg Leu His Gly Ile Pro His Val Ala Thr	
115 120 125	
gcg cat agc ctg gag ccg gac cgg ccc tgg aag cgc gag caa ctc ggc	432
Ala His Ser Leu Glu Pro Asp Arg Pro Trp Lys Arg Glu Gln Leu Gly	

-continued

130	135	140	
ggc ggc tac gac gtg agc tcc tgg tcg gag aaa aac gcg atg gag tac Gly Gly Tyr Asp Val Ser Ser Trp Ser Glu Lys Asn Ala Met Glu Tyr 145 150 155 160			480
gcg gac gcc gtg atc gcc gtc agt gcc cgg atg aaa gac tcc atc ctg Ala Asp Ala Val Ile Ala Val Ser Ala Arg Met Lys Asp Ser Ile Leu 165 170 175			528
gcg gct tat ccg cgc atc gag ccc gat aat gtg cgc gtg gtg ctg aac Ala Ala Tyr Pro Arg Ile Glu Pro Asp Asn Val Arg Val Val Leu Asn 180 185 190			576
ggc atc gac acc gag ctc tgg cag ccg agg ccg acc ttc gac gac gcc Gly Ile Asp Thr Glu Leu Trp Gln Pro Arg Pro Thr Phe Asp Asp Ala 195 200 205			624
gag gat tcc gtg ctg cgc agc ctg gcc gtc gac ccg caa cgg ccc atc Glu Asp Ser Val Leu Arg Ser Leu Gly Val Asp Pro Gln Arg Pro Ile 210 215 220			672
gtc gcg ttt gtc gga cgg att acg cgg cag aaa gcc gtg gag cac ctc Val Ala Phe Val Gly Arg Ile Thr Arg Gln Lys Gly Val Glu His Leu 225 230 235 240			720
atc aaa gcc gcc gcc ctg ttc gac gag tcc gtc cag ctc gtc ctc tgc Ile Lys Ala Ala Ala Leu Phe Asp Glu Ser Val Gln Leu Val Leu Cys 245 250 255			768
gcg ggt gcc ccc gac acc ccg gag atc gcg gct cgg acc acg gcg ctg Ala Gly Ala Pro Asp Thr Pro Glu Ile Ala Ala Arg Thr Thr Ala Leu 260 265 270			816
gtc gag gaa ctc caa gcg aag cgc gag gcc atc ttc tgg gtc cag gat Val Glu Glu Leu Gln Ala Lys Arg Glu Gly Ile Phe Trp Val Gln Asp 275 280 285			864
atg ctg ggg aag gat aag atc cag gag atc ctc acc gcc gct gac acc Met Leu Gly Lys Asp Lys Ile Gln Glu Ile Leu Thr Ala Ala Asp Thr 290 295 300			912
ttc gtg tgc ccg tcg atc tat gag ccc ctg gcc atc gtc aac ctc gaa Phe Val Cys Pro Ser Ile Tyr Glu Pro Leu Gly Ile Val Asn Leu Glu 305 310 315 320			960
gcc atg gcg tgc aat acc gcc gtg gtc gcg agc gac gtc gcc gcc atc Ala Met Ala Cys Asn Thr Ala Val Val Ala Ser Asp Val Gly Gly Ile 325 330 335			1008
cca gag gtc gtc gtg gac gcc acc acg gcc gca ctg gtg cat tac gat Pro Glu Val Val Val Asp Gly Thr Thr Gly Ala Leu Val His Tyr Asp 340 345 350			1056
gag aac gat gtg gaa acg ttc gag cgc gac att gcc gaa gcc gtg aac Glu Asn Asp Val Glu Thr Phe Glu Arg Asp Ile Ala Glu Ala Val Asn 355 360 365			1104
aag atg gtc gcg gat cgc gag act gcc gcg aag ttc ggt ctt gca gcc Lys Met Val Ala Asp Arg Glu Thr Ala Ala Lys Phe Gly Leu Ala Gly 370 375 380			1152
egg gag cgg gcg atc aat gac ttc agc tgg gcc acc atc gcc cag cag Arg Glu Arg Ala Ile Asn Asp Phe Ser Trp Ala Thr Ile Ala Gln Gln 385 390 395 400			1200
acc atc gac gtc tat aag tcg ctg atg Thr Ile Asp Val Tyr Lys Ser Leu Met 405			1227

<210> SEQ ID NO 34

<211> LENGTH: 409

<212> TYPE: PRT

<213> ORGANISM: Artificial sequence

<220> FEATURE:

-continued

<223> OTHER INFORMATION: Synthetic Construct

<400> SEQUENCE: 34

Met Pro Pro Phe Arg Tyr Arg Cys Ala Thr Val Phe Arg Trp Leu Ile
1 5 10 15

Phe Glu Ile Met Arg Val Gly Met Met Thr Arg Glu Tyr Pro Pro Glu
20 25 30

Val Tyr Gly Gly Ala Gly Val His Val Thr Glu Leu Thr Arg Phe Met
35 40 45

Arg Glu Ile Ala Glu Val Asp Val His Cys Met Gly Ala Pro Arg Asp
50 55 60

Met Glu Gly Val Phe Val His Gly Val Asp Pro Ala Leu Glu Ser Ala
65 70 75 80

Asn Pro Ala Ile Lys Thr Leu Ser Thr Gly Leu Arg Met Ala Glu Ala
85 90 95

Ala Asn Asn Val Asp Val Val His Ser His Thr Trp Tyr Ala Gly Leu
100 105 110

Gly Gly His Leu Ala Ala Arg Leu His Gly Ile Pro His Val Ala Thr
115 120 125

Ala His Ser Leu Glu Pro Asp Arg Pro Trp Lys Arg Glu Gln Leu Gly
130 135 140

Gly Gly Tyr Asp Val Ser Ser Trp Ser Glu Lys Asn Ala Met Glu Tyr
145 150 155 160

Ala Asp Ala Val Ile Ala Val Ser Ala Arg Met Lys Asp Ser Ile Leu
165 170 175

Ala Ala Tyr Pro Arg Ile Glu Pro Asp Asn Val Arg Val Val Leu Asn
180 185 190

Gly Ile Asp Thr Glu Leu Trp Gln Pro Arg Pro Thr Phe Asp Asp Ala
195 200 205

Glu Asp Ser Val Leu Arg Ser Leu Gly Val Asp Pro Gln Arg Pro Ile
210 215 220

Val Ala Phe Val Gly Arg Ile Thr Arg Gln Lys Gly Val Glu His Leu
225 230 235 240

Ile Lys Ala Ala Ala Leu Phe Asp Glu Ser Val Gln Leu Val Leu Cys
245 250 255

Ala Gly Ala Pro Asp Thr Pro Glu Ile Ala Ala Arg Thr Thr Ala Leu
260 265 270

Val Glu Glu Leu Gln Ala Lys Arg Glu Gly Ile Phe Trp Val Gln Asp
275 280 285

Met Leu Gly Lys Asp Lys Ile Gln Glu Ile Leu Thr Ala Ala Asp Thr
290 295 300

Phe Val Cys Pro Ser Ile Tyr Glu Pro Leu Gly Ile Val Asn Leu Glu
305 310 315 320

Ala Met Ala Cys Asn Thr Ala Val Val Ala Ser Asp Val Gly Gly Ile
325 330 335

Pro Glu Val Val Val Asp Gly Thr Thr Gly Ala Leu Val His Tyr Asp
340 345 350

Glu Asn Asp Val Glu Thr Phe Glu Arg Asp Ile Ala Glu Ala Val Asn
355 360 365

Lys Met Val Ala Asp Arg Glu Thr Ala Ala Lys Phe Gly Leu Ala Gly
370 375 380

-continued

Arg Glu Arg Ala Ile Asn Asp Phe Ser Trp Ala Thr Ile Ala Gln Gln
385 390 395 400

Thr Ile Asp Val Tyr Lys Ser Leu Met
405

<210> SEQ ID NO 35
 <211> LENGTH: 2184
 <212> TYPE: DNA
 <213> ORGANISM: Artificial sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Synthetic construct
 <220> FEATURE:
 <221> NAME/KEY: CDS
 <222> LOCATION: (1)..(2184)
 <223> OTHER INFORMATION: E. coli 1,4-alpha-glucan branching enzyme
 (Acc.No. YP 492001.1) (codon optimized)

<400> SEQUENCE: 35

atg tcc gac cgc att gat agg gac gtc ata aat gca ctg atc gct ggc	48
Met Ser Asp Arg Ile Asp Arg Asp Val Ile Asn Ala Leu Ile Ala Gly	
1 5 10 15	
cac ttt gct gac ccg ttc tcc gtt ctg ggc atg cat aag acc acc gcc	96
His Phe Ala Asp Pro Phe Ser Val Leu Gly Met His Lys Thr Thr Ala	
20 25 30	
ggt ctg gag gtc cgc gcg ctg ctg ccc gac gcg acc gac gtc tgg gtc	144
Gly Leu Glu Val Arg Ala Leu Leu Pro Asp Ala Thr Asp Val Trp Val	
35 40 45	
atc gag ccc aag act ggc cgc aaa ctg gcg aaa ctt gag tgc ctc gac	192
Ile Glu Pro Lys Thr Gly Arg Lys Leu Ala Lys Leu Glu Cys Leu Asp	
50 55 60	
agc cgg gga ttc ttt agc ggc gtg atc ccg cgg cgg aag aac ttc ttt	240
Ser Arg Gly Phe Phe Ser Gly Val Ile Pro Arg Arg Lys Asn Phe Phe	
65 70 75 80	
agg tat caa ctc gcc gtc gtg tgg cat ggg cag cag aac ctg atc gat	288
Arg Tyr Gln Leu Ala Val Val Trp His Gly Gln Gln Asn Leu Ile Asp	
85 90 95	
gat ccc tac agg ttc ggt ccc ttg atc caa gag atg gat gcg tgg ctg	336
Asp Pro Tyr Arg Phe Gly Pro Leu Ile Gln Glu Met Asp Ala Trp Leu	
100 105 110	
ctc tcc gag ggc acc cac ctc cgc ccg tac gag act ctc ggg gca cat	384
Leu Ser Glu Gly Thr His Leu Arg Pro Tyr Glu Thr Leu Gly Ala His	
115 120 125	
gcg gac acg atg gac ggc gtg acg ggc acc cgc ttc tcg gtc tgg gct	432
Ala Asp Thr Met Asp Gly Val Thr Gly Thr Arg Phe Ser Val Trp Ala	
130 135 140	
ccg aat gca cgc cgg gtg tcc gtg gtc gga cag ttc aat tac tgg gat	480
Pro Asn Ala Arg Arg Val Ser Val Val Gly Gln Phe Asn Tyr Trp Asp	
145 150 155 160	
ggt cgg cgc cac ccc atg cgg ctg cgg aag gaa tcg ggt atc tgg gag	528
Gly Arg Arg His Pro Met Arg Leu Arg Lys Glu Ser Gly Ile Trp Glu	
165 170 175	
ttg ttc atc cca ggc gcc cat aac ggg cag ctc tac aaa tac gaa atg	576
Leu Phe Ile Pro Gly Ala His Asn Gly Gln Leu Tyr Lys Tyr Glu Met	
180 185 190	
atc gat gcg aat ggc aac ctc cgg ctg aaa agt gac ccg tat gcc ttc	624
Ile Asp Ala Asn Gly Asn Leu Arg Leu Lys Ser Asp Pro Tyr Ala Phe	
195 200 205	
gaa gcg cag atg cgc ccc gaa acg gcg tcc ctg atc tgc ggc ctc cct	672
Glu Ala Gln Met Arg Pro Glu Thr Ala Ser Leu Ile Cys Gly Leu Pro	
210 215 220	

-continued

gaa aaa gtc gtc cag acc gag gaa cgc aag aag gcc aac caa ttc gac	720
Glu Lys Val Val Gln Thr Glu Glu Arg Lys Lys Ala Asn Gln Phe Asp	
225 230 235 240	
gcg ccg atc tcg atc tat gag gtc cac ctg ggc tcg tgg cgc agg cac	768
Ala Pro Ile Ser Ile Tyr Glu Val His Leu Gly Ser Trp Arg Arg His	
245 250 255	
acc gac aac aat ttc tgg ctc tcg tac cgc gag ctg gcg gac caa ctc	816
Thr Asp Asn Asn Phe Trp Leu Ser Tyr Arg Glu Leu Ala Asp Gln Leu	
260 265 270	
gtg ccg tat gct aag tgg atg gga ttc acg cat ttg gaa ctg ctg ccg	864
Val Pro Tyr Ala Lys Trp Met Gly Phe Thr His Leu Glu Leu Leu Pro	
275 280 285	
atc aac gaa cat ccc ttc gac ggc agc tgg ggc tat cag ccg acc ggc	912
Ile Asn Glu His Pro Phe Asp Gly Ser Trp Gly Tyr Gln Pro Thr Gly	
290 295 300	
ctc tac gcc ccg act cgc cgg ttc ggc acg cgg gat gac ttc cgg tac	960
Leu Tyr Ala Pro Thr Arg Arg Phe Gly Thr Arg Asp Asp Phe Arg Tyr	
305 310 315 320	
ttc atc gat gcc gcg cat gcc gcc ggc ctc aac gtc atc ctg gac tgg	1008
Phe Ile Asp Ala Ala His Ala Ala Gly Leu Asn Val Ile Leu Asp Trp	
325 330 335	
gtg ccc ggt cac ttt ccc acc gac gac ttc gcg ctg gcc gag ttc gac	1056
Val Pro Gly His Phe Pro Thr Asp Asp Phe Ala Leu Ala Glu Phe Asp	
340 345 350	
ggc acc aac ctc tac gag cat agt gat ccg cgc gag gcc tac cat cag	1104
Gly Thr Asn Leu Tyr Glu His Ser Asp Pro Arg Glu Gly Tyr His Gln	
355 360 365	
gac tgg aac acg ctc atc tac aat tac ggt cgc cgc gag gtc agc aac	1152
Asp Trp Asn Thr Leu Ile Tyr Asn Tyr Gly Arg Arg Glu Val Ser Asn	
370 375 380	
ttc ctg gtt ggg aat gcg ctg tat tgg att gag cgc ttc gcc ata gac	1200
Phe Leu Val Gly Asn Ala Leu Tyr Trp Ile Glu Arg Phe Gly Ile Asp	
385 390 395 400	
gcc ctg cgc gtc gac gcc gtg gca tcc atg atc tac cgc gat tat tcc	1248
Ala Leu Arg Val Asp Ala Val Ala Ser Met Ile Tyr Arg Asp Tyr Ser	
405 410 415	
cgc aaa gag ggc gag tgg atc ccc aat gag ttc ggt gcc cgc gag aac	1296
Arg Lys Glu Gly Glu Trp Ile Pro Asn Glu Phe Gly Gly Arg Glu Asn	
420 425 430	
ctt gag gcc att gag ttc ctt agg aat acg aac cgg atc ctg ggg gaa	1344
Leu Glu Ala Ile Glu Phe Leu Arg Asn Thr Asn Arg Ile Leu Gly Glu	
435 440 445	
caa gtg tcc ggg gct gtc acc atg gca gag gag agc acc gat ttt ccc	1392
Gln Val Ser Gly Ala Val Thr Met Ala Glu Glu Ser Thr Asp Phe Pro	
450 455 460	
ggc gtg tcg cgc ccg caa gac atg ggt gcc ctg gcc ttc tgg tac aag	1440
Gly Val Ser Arg Pro Gln Asp Met Gly Gly Leu Gly Phe Trp Tyr Lys	
465 470 475 480	
tgg aat ctg gcc tgg atg cac gac acc ctc gac tac atg aag ctt gat	1488
Trp Asn Leu Gly Trp Met His Asp Thr Leu Asp Tyr Met Lys Leu Asp	
485 490 495	
ccg gtc tat cgc cag tat cac cat gac aag ctc acg ttc gcc atc ctg	1536
Pro Val Tyr Arg Gln Tyr His His Asp Lys Leu Thr Phe Gly Ile Leu	
500 505 510	
tat aac tat acc gag aat ttc gtg ctc ccg ttg agc cat gac gaa gtt	1584
Tyr Asn Tyr Thr Glu Asn Phe Val Leu Pro Leu Ser His Asp Glu Val	
515 520 525	

-continued

gtc cat ggc aag aag agt att ctg gac cgg atg cca ggc gac gcg tgg	1632
Val His Gly Lys Lys Ser Ile Leu Asp Arg Met Pro Gly Asp Ala Trp	
530 535 540	
cag aaa ttc gcg aat ctc cgc gcc tat tat ggc tgg atg tgg gcg ttc	1680
Gln Lys Phe Ala Asn Leu Arg Ala Tyr Tyr Gly Trp Met Trp Ala Phe	
545 550 555 560	
ccg ggc aaa aag ctc ctg ttc atg gga aat gag ttc gcc cag ggc cgg	1728
Pro Gly Lys Lys Leu Leu Phe Met Gly Asn Glu Phe Ala Gln Gly Arg	
565 570 575	
gag tgg aac cat gac gcg agc ctc gac tgg cat ctc ctt gag ggc gga	1776
Glu Trp Asn His Asp Ala Ser Leu Asp Trp His Leu Leu Glu Gly Gly	
580 585 590	
gac aac tgg cac cac ggc gtg cag cgc ctc gtg cgg gac ctc aac ctg	1824
Asp Asn Trp His His Gly Val Gln Arg Leu Val Arg Asp Leu Asn Leu	
595 600 605	
acc tac cgc cat cat aaa gcc atg cac gag ctg gat ttc gac ccg tac	1872
Thr Tyr Arg His His Lys Ala Met His Glu Leu Asp Phe Asp Pro Tyr	
610 615 620	
ggc ttc gag tgg ctc gtc gtc gac gat aag gag cgc tcg gtc ctc att	1920
Gly Phe Glu Trp Leu Val Val Asp Asp Lys Glu Arg Ser Val Leu Ile	
625 630 635 640	
ttc gtg cgc agg gac aag gaa ggc aac gag atc atc gtg gcg agc aac	1968
Phe Val Arg Arg Asp Lys Glu Gly Asn Glu Ile Ile Val Ala Ser Asn	
645 650 655	
ttc acc ccg gtc ccg cgg cac gac tac cgc ttc ggc atc aat cag ccg	2016
Phe Thr Pro Val Pro Arg His Asp Tyr Arg Phe Gly Ile Asn Gln Pro	
660 665 670	
ggc aag tgg cgc gag atc ctg aac acg gac tcg atg cat tat cat ggt	2064
Gly Lys Trp Arg Glu Ile Leu Asn Thr Asp Ser Met His Tyr His Gly	
675 680 685	
tcg aac gcc ggg aat ggc ggc acc gtg cac tcg gac gag atc gcc tcc	2112
Ser Asn Ala Gly Asn Gly Gly Thr Val His Ser Asp Glu Ile Ala Ser	
690 695 700	
cat ggc cgc cag cat agc ttg tcc ctg acc ctg ccc cct ctc gcg acc	2160
His Gly Arg Gln His Ser Leu Ser Leu Thr Leu Pro Pro Leu Ala Thr	
705 710 715 720	
atc tgg ctg gtg cgc gag gcc gag	2184
Ile Trp Leu Val Arg Glu Ala Glu	
725	
<210> SEQ ID NO 36	
<211> LENGTH: 728	
<212> TYPE: PRT	
<213> ORGANISM: Artificial sequence	
<220> FEATURE:	
<223> OTHER INFORMATION: Synthetic Construct	
<400> SEQUENCE: 36	
Met Ser Asp Arg Ile Asp Arg Asp Val Ile Asn Ala Leu Ile Ala Gly	
1 5 10 15	
His Phe Ala Asp Pro Phe Ser Val Leu Gly Met His Lys Thr Thr Ala	
20 25 30	
Gly Leu Glu Val Arg Ala Leu Leu Pro Asp Ala Thr Asp Val Trp Val	
35 40 45	
Ile Glu Pro Lys Thr Gly Arg Lys Leu Ala Lys Leu Glu Cys Leu Asp	
50 55 60	
Ser Arg Gly Phe Phe Ser Gly Val Ile Pro Arg Arg Lys Asn Phe Phe	

-continued

65	70					75					80				
Arg Tyr Gln Leu Ala Val Val Trp His Gly Gln Gln Asn Leu Ile Asp	85					90					95				
Asp Pro Tyr Arg Phe Gly Pro Leu Ile Gln Glu Met Asp Ala Trp Leu	100					105					110				
Leu Ser Glu Gly Thr His Leu Arg Pro Tyr Glu Thr Leu Gly Ala His	115					120					125				
Ala Asp Thr Met Asp Gly Val Thr Gly Thr Arg Phe Ser Val Trp Ala	130					135					140				
Pro Asn Ala Arg Arg Val Ser Val Val Gly Gln Phe Asn Tyr Trp Asp	145					150					155				
Gly Arg Arg His Pro Met Arg Leu Arg Lys Glu Ser Gly Ile Trp Glu	165					170					175				
Leu Phe Ile Pro Gly Ala His Asn Gly Gln Leu Tyr Lys Tyr Glu Met	180					185					190				
Ile Asp Ala Asn Gly Asn Leu Arg Leu Lys Ser Asp Pro Tyr Ala Phe	195					200					205				
Glu Ala Gln Met Arg Pro Glu Thr Ala Ser Leu Ile Cys Gly Leu Pro	210					215					220				
Glu Lys Val Val Gln Thr Glu Glu Arg Lys Lys Ala Asn Gln Phe Asp	225					230					235				
Ala Pro Ile Ser Ile Tyr Glu Val His Leu Gly Ser Trp Arg Arg His	245					250					255				
Thr Asp Asn Asn Phe Trp Leu Ser Tyr Arg Glu Leu Ala Asp Gln Leu	260					265					270				
Val Pro Tyr Ala Lys Trp Met Gly Phe Thr His Leu Glu Leu Leu Pro	275					280					285				
Ile Asn Glu His Pro Phe Asp Gly Ser Trp Gly Tyr Gln Pro Thr Gly	290					295					300				
Leu Tyr Ala Pro Thr Arg Arg Phe Gly Thr Arg Asp Asp Phe Arg Tyr	305					310					315				
Phe Ile Asp Ala Ala His Ala Ala Gly Leu Asn Val Ile Leu Asp Trp	325					330					335				
Val Pro Gly His Phe Pro Thr Asp Asp Phe Ala Leu Ala Glu Phe Asp	340					345					350				
Gly Thr Asn Leu Tyr Glu His Ser Asp Pro Arg Glu Gly Tyr His Gln	355					360					365				
Asp Trp Asn Thr Leu Ile Tyr Asn Tyr Gly Arg Arg Glu Val Ser Asn	370					375					380				
Phe Leu Val Gly Asn Ala Leu Tyr Trp Ile Glu Arg Phe Gly Ile Asp	385					390					395				
Ala Leu Arg Val Asp Ala Val Ala Ser Met Ile Tyr Arg Asp Tyr Ser	405					410					415				
Arg Lys Glu Gly Glu Trp Ile Pro Asn Glu Phe Gly Gly Arg Glu Asn	420					425					430				
Leu Glu Ala Ile Glu Phe Leu Arg Asn Thr Asn Arg Ile Leu Gly Glu	435					440					445				
Gln Val Ser Gly Ala Val Thr Met Ala Glu Glu Ser Thr Asp Phe Pro	450					455					460				
Gly Val Ser Arg Pro Gln Asp Met Gly Gly Leu Gly Phe Trp Tyr Lys	465					470					475				
											480				

-continued

Trp Asn Leu Gly Trp Met His Asp Thr Leu Asp Tyr Met Lys Leu Asp
 485 490 495

Pro Val Tyr Arg Gln Tyr His His Asp Lys Leu Thr Phe Gly Ile Leu
 500 505 510

Tyr Asn Tyr Thr Glu Asn Phe Val Leu Pro Leu Ser His Asp Glu Val
 515 520 525

Val His Gly Lys Lys Ser Ile Leu Asp Arg Met Pro Gly Asp Ala Trp
 530 535 540

Gln Lys Phe Ala Asn Leu Arg Ala Tyr Tyr Gly Trp Met Trp Ala Phe
 545 550 555 560

Pro Gly Lys Lys Leu Leu Phe Met Gly Asn Glu Phe Ala Gln Gly Arg
 565 570 575

Glu Trp Asn His Asp Ala Ser Leu Asp Trp His Leu Leu Glu Gly Gly
 580 585 590

Asp Asn Trp His His Gly Val Gln Arg Leu Val Arg Asp Leu Asn Leu
 595 600 605

Thr Tyr Arg His His Lys Ala Met His Glu Leu Asp Phe Asp Pro Tyr
 610 615 620

Gly Phe Glu Trp Leu Val Val Asp Asp Lys Glu Arg Ser Val Leu Ile
 625 630 635 640

Phe Val Arg Arg Asp Lys Glu Gly Asn Glu Ile Ile Val Ala Ser Asn
 645 650 655

Phe Thr Pro Val Pro Arg His Asp Tyr Arg Phe Gly Ile Asn Gln Pro
 660 665 670

Gly Lys Trp Arg Glu Ile Leu Asn Thr Asp Ser Met His Tyr His Gly
 675 680 685

Ser Asn Ala Gly Asn Gly Gly Thr Val His Ser Asp Glu Ile Ala Ser
 690 695 700

His Gly Arg Gln His Ser Leu Ser Leu Thr Leu Pro Pro Leu Ala Thr
 705 710 715 720

Ile Trp Leu Val Arg Glu Ala Glu
 725

<210> SEQ ID NO 37
 <211> LENGTH: 2193
 <212> TYPE: DNA
 <213> ORGANISM: Artificial sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Synthetic construct
 <220> FEATURE:
 <221> NAME/KEY: CDS
 <222> LOCATION: (1)..(2193)
 <223> OTHER INFORMATION: Corynebacterium glutamicum (ATCC 13032)
 Glycogen branching enzyme (codon optimized)

<400> SEQUENCE: 37

atg acc gtc gac ccc gcg tcc cat atc acg atc ccc gaa gcc gac ctc	48
Met Thr Val Asp Pro Ala Ser His Ile Thr Ile Pro Glu Ala Asp Leu	
1 5 10 15	
gca cgc ctc cgc cac tgc aac cat cac gac ccc cat gga ttc tac ggg	96
Ala Arg Leu Arg His Cys Asn His His Asp Pro His Gly Phe Tyr Gly	
20 25 30	
tgg cat gaa act gag gcc ggg tcc gtc att cgc acc cgc cag gtc ggc	144
Trp His Glu Thr Glu Ala Gly Ser Val Ile Arg Thr Arg Gln Val Gly	
35 40 45	

-continued

gcg acc cag gtg aat ctg ctc att gat gac acg agc cat gtc atg acg Ala Thr Gln Val Asn Leu Leu Ile Asp Asp Thr Ser His Val Met Thr 50 55 60	192
ccc atc ggc gac gac atc ttc gcg atc gac ctc ggg cat cgg gag cgc Pro Ile Gly Asp Asp Ile Phe Ala Ile Asp Leu Gly His Arg Glu Arg 65 70 75 80	240
gct gat tac cgc ctt gag gtc acg tgg ccg gac caa gag cca cag gtg Ala Asp Tyr Arg Glu Val Thr Trp Pro Asp Gln Glu Pro Gln Val 85 90 95	288
aaa gct gac ccg tac tat ttc ctc ccc acg gtc ggt gag atg gac atc Lys Ala Asp Pro Tyr Tyr Phe Leu Pro Thr Val Gly Glu Met Asp Ile 100 105 110	336
tat ctg ttc tcc gag ggt cgg cac gaa ccg ctc tgg gag atc ctc ggc Tyr Leu Phe Ser Glu Gly Arg His Glu Arg Leu Trp Glu Ile Leu Gly 115 120 125	384
gcc aat atc aag acc tat cag acc gcg ctg ggc acc gtc cgc ggc acg Ala Asn Ile Lys Thr Tyr Gln Thr Ala Leu Gly Thr Val Arg Gly Thr 130 135 140	432
gcg ttc acc gtc tgg gca ccc aac gcg atc ggc tgc gcg gtc gtg ggg Ala Phe Thr Val Trp Ala Pro Asn Ala Ile Gly Cys Ala Val Val Gly 145 150 155 160	480
ggc ttc aac ggt tgg aac gcc agt caa cac ccg atg ccg agc atg ggc Gly Phe Asn Gly Trp Asn Ala Ser Gln His Pro Met Arg Ser Met Gly 165 170 175	528
gga tcc ggc ctt tgg gag ctg ttc atc ccg ggc atc gag gag ggc gag Gly Ser Gly Leu Trp Glu Leu Phe Ile Pro Gly Ile Glu Glu Gly Glu 180 185 190	576
gtg tac aag ttt gcg gtg cag acc cgc gaa ggc cag cgc agg gat aag Val Tyr Lys Phe Ala Val Gln Thr Arg Glu Gly Gln Arg Arg Asp Lys 195 200 205	624
gcc gac ccc atg gcg ccg cgc gcc gag ctg gca cca gcc acc ggc agc Ala Asp Pro Met Ala Arg Arg Ala Glu Leu Ala Pro Ala Thr Gly Ser 210 215 220	672
ata gtt gcc tcg tcc gag tac cag tgg cag gac tcc gag tgg ctg cgc Ile Val Ala Ser Ser Glu Tyr Gln Trp Gln Asp Ser Glu Trp Leu Arg 225 230 235 240	720
gag ccg agc cag act gac ctc gcc agc aag cct atg tcc gtg tat gag Glu Arg Ser Gln Thr Asp Leu Ala Ser Lys Pro Met Ser Val Tyr Glu 245 250 255	768
gtg cat ctg ggt tcg tgg cgc tgg ggc aag aac tac gaa gat ctg gcg Val His Leu Gly Ser Trp Arg Trp Gly Lys Asn Tyr Glu Asp Leu Ala 260 265 270	816
acc gag ctc gtc gat tat gtt gcg gac ctg ggt tat acg cat gtc gag Thr Glu Leu Val Asp Tyr Val Ala Asp Leu Gly Tyr Thr His Val Glu 275 280 285	864
ttc ctg ccg gtg gcg gag cac ccg ttc ggc ggc tcg tgg ggc tac caa Phe Leu Pro Val Ala Glu His Pro Phe Gly Gly Ser Trp Gly Tyr Gln 290 295 300	912
gtc acg ggt tat tac gcg ccg acc agc cgc tgg ggc acc ccg gac cag Val Thr Gly Tyr Tyr Ala Pro Thr Ser Arg Trp Gly Thr Pro Asp Gln 305 310 315 320	960
ttc cgc gcg ttg gtc gac gcg ttc cac gcc agg gga atc ggc gtc atc Phe Arg Ala Leu Val Asp Ala Phe His Ala Arg Gly Ile Gly Val Ile 325 330 335	1008
atg gac tgg gtt ccc gca cat ttc ccg aaa gat gat tgg gcg ctg gcg Met Asp Trp Val Pro Ala His Phe Pro Lys Asp Asp Trp Ala Leu Ala 340 345 350	1056

-continued

cgg ttc gac ggc gag gcc ctc tat gag cac ccg gac tgg cgg cgg ggt	1104
Arg Phe Asp Gly Glu Ala Leu Tyr Glu His Pro Asp Trp Arg Arg Gly	
355 360 365	
gag cag aaa gat tgg ggg acc ctc gtg ttc gat ttt ggc cgg aac gaa	1152
Glu Gln Lys Asp Trp Gly Thr Leu Val Phe Asp Phe Gly Arg Asn Glu	
370 375 380	
gtc cgc aac ttc ttg gtc gcg aac gcg ctc tat tgg att gaa gag ttc	1200
Val Arg Asn Phe Leu Val Ala Asn Ala Leu Tyr Trp Ile Glu Glu Phe	
385 390 395 400	
cat att gac ggc ctc cgg gtc gac gcc gtc gcc tcg atg ctc tac ctg	1248
His Ile Asp Gly Leu Arg Val Asp Ala Val Ala Ser Met Leu Tyr Leu	
405 410 415	
gac tac tcg cgc gag cac ggc gag tgg gag ccg aac atc tat ggc gga	1296
Asp Tyr Ser Arg Glu His Gly Glu Trp Glu Pro Asn Ile Tyr Gly Gly	
420 425 430	
cgg gag aat ctc gaa gcc gtc cag ttc ctg caa gag atg aac gca acc	1344
Arg Glu Asn Leu Glu Ala Val Gln Phe Leu Gln Glu Met Asn Ala Thr	
435 440 445	
gtg ctc cgc ctg cat cct ggt gcg ctg acc atc gca gag gaa tcg acc	1392
Val Leu Arg Leu His Pro Gly Ala Leu Thr Ile Ala Glu Glu Ser Thr	
450 455 460	
tcg tgg ccg ggt gtg acc gct ccg acc tgg gac ggt ggc ctt ggt ttc	1440
Ser Trp Pro Gly Val Thr Ala Pro Thr Trp Asp Gly Gly Leu Gly Phe	
465 470 475 480	
agc ctg aag tgg aat atg ggc tgg atg cat gac acc ctt gag tat ttc	1488
Ser Leu Lys Trp Asn Met Gly Trp Met His Asp Thr Leu Glu Tyr Phe	
485 490 495	
tcg aag aat ccg gtg cac agg gcc ttc cac cat tcc gag ctg acg ttc	1536
Ser Lys Asn Pro Val His Arg Ala Phe His His Ser Glu Leu Thr Phe	
500 505 510	
tcg ctt gtc tac gcc ttc agt gag cgc ttc gtg ctg ccg atc tcc cat	1584
Ser Leu Val Tyr Ala Phe Ser Glu Arg Phe Val Leu Pro Ile Ser His	
515 520 525	
gac gaa gtc gtc cac ggc aag ggc tcc ctg tgg gac ccg atg ccg ggt	1632
Asp Glu Val Val His Gly Lys Ser Leu Trp Asp Arg Met Pro Gly	
530 535 540	
gac acc tgg aat aaa gcc gca ggc ctc cgc acg ttc ctg gcg tac atg	1680
Asp Thr Trp Asn Lys Ala Ala Gly Leu Arg Thr Phe Leu Ala Tyr Met	
545 550 555 560	
tgg agc cac ccg ggc aag aaa ctg ctg ttc atg ggc caa gag ttc ggc	1728
Trp Ser His Pro Gly Lys Lys Leu Leu Phe Met Gly Gln Glu Phe Gly	
565 570 575	
cag cgc gag gag tgg gcc gag ggc cag ggc ctg ccc tgg gac atc gtg	1776
Gln Arg Glu Glu Trp Ala Glu Gly Gln Gly Leu Pro Trp Asp Ile Val	
580 585 590	
gac ggg tgg cag ggc gag tat cat gaa gcg atc cgc acc ctc acg cgc	1824
Asp Gly Trp Gln Gly Glu Tyr His Glu Ala Ile Arg Thr Leu Thr Arg	
595 600 605	
agc ctg aac ggc gtg tac agc gat agt ccg gcg ctc cat acc caa gat	1872
Ser Leu Asn Gly Val Tyr Ser Asp Ser Pro Ala Leu His Thr Gln Asp	
610 615 620	
ttc acc ggc gag ggc ttc acg tgg aac aaa ggt gac gac gcc acc aat	1920
Phe Thr Gly Glu Gly Phe Thr Trp Asn Lys Gly Asp Asp Ala Thr Asn	
625 630 635 640	
aac atc ctc gcc ttc acc cgc ttt ggc tcc gac ggc tcg cag atg ctc	1968
Asn Ile Leu Ala Phe Thr Arg Phe Gly Ser Asp Gly Ser Gln Met Leu	
645 650 655	

-continued

```

tgc gtg ttc aac ctc tcg ggc acc tcg cag ccg gag tac cag ctc ggg 2016
Cys Val Phe Asn Leu Ser Gly Thr Ser Gln Pro Glu Tyr Gln Leu Gly
        660                665                670

gtg gcc gct ggg ggc gag tgg aag ctt gtg ctg aat acc gac gac gcc 2064
Val Ala Ala Gly Gly Glu Trp Lys Leu Val Leu Asn Thr Asp Asp Ala
        675                680                685

gag ttt ttg gga gcc gaa aac gat atc gcg acg agc gtg caa gct gct 2112
Glu Phe Leu Gly Ala Glu Asn Asp Ile Ala Thr Ser Val Gln Ala Ala
        690                695                700

gcc act ccc agg gac aat ttc gcc tat tcc ctg agc ctg cat gtc ccc 2160
Ala Thr Pro Arg Asp Asn Phe Ala Tyr Ser Leu Ser Leu His Val Pro
        705                710                715                720

gcc atg tcg gcg cag ttc tac tcg ctc cag aag 2193
Ala Met Ser Ala Gln Phe Tyr Ser Leu Gln Lys
        725                730
    
```

```

<210> SEQ ID NO 38
<211> LENGTH: 731
<212> TYPE: PRT
<213> ORGANISM: Artificial sequence
<220> FEATURE:
<223> OTHER INFORMATION: Synthetic Construct
    
```

<400> SEQUENCE: 38

```

Met Thr Val Asp Pro Ala Ser His Ile Thr Ile Pro Glu Ala Asp Leu
1          5          10          15

Ala Arg Leu Arg His Cys Asn His His Asp Pro His Gly Phe Tyr Gly
20          25          30

Trp His Glu Thr Glu Ala Gly Ser Val Ile Arg Thr Arg Gln Val Gly
35          40          45

Ala Thr Gln Val Asn Leu Leu Ile Asp Asp Thr Ser His Val Met Thr
50          55          60

Pro Ile Gly Asp Asp Ile Phe Ala Ile Asp Leu Gly His Arg Glu Arg
65          70          75          80

Ala Asp Tyr Arg Leu Glu Val Thr Trp Pro Asp Gln Glu Pro Gln Val
85          90          95

Lys Ala Asp Pro Tyr Tyr Phe Leu Pro Thr Val Gly Glu Met Asp Ile
100         105         110

Tyr Leu Phe Ser Glu Gly Arg His Glu Arg Leu Trp Glu Ile Leu Gly
115         120         125

Ala Asn Ile Lys Thr Tyr Gln Thr Ala Leu Gly Thr Val Arg Gly Thr
130         135         140

Ala Phe Thr Val Trp Ala Pro Asn Ala Ile Gly Cys Ala Val Val Gly
145         150         155         160

Gly Phe Asn Gly Trp Asn Ala Ser Gln His Pro Met Arg Ser Met Gly
165         170         175

Gly Ser Gly Leu Trp Glu Leu Phe Ile Pro Gly Ile Glu Glu Gly Glu
180         185         190

Val Tyr Lys Phe Ala Val Gln Thr Arg Glu Gly Gln Arg Arg Asp Lys
195         200         205

Ala Asp Pro Met Ala Arg Arg Ala Glu Leu Ala Pro Ala Thr Gly Ser
210         215         220

Ile Val Ala Ser Ser Glu Tyr Gln Trp Gln Asp Ser Glu Trp Leu Arg
225         230         235         240

Glu Arg Ser Gln Thr Asp Leu Ala Ser Lys Pro Met Ser Val Tyr Glu
    
```

-continued

245					250					255					
Val	His	Leu	Gly	Ser	Trp	Arg	Trp	Gly	Lys	Asn	Tyr	Glu	Asp	Leu	Ala
		260						265					270		
Thr	Glu	Leu	Val	Asp	Tyr	Val	Ala	Asp	Leu	Gly	Tyr	Thr	His	Val	Glu
		275					280						285		
Phe	Leu	Pro	Val	Ala	Glu	His	Pro	Phe	Gly	Gly	Ser	Trp	Gly	Tyr	Gln
	290					295					300				
Val	Thr	Gly	Tyr	Tyr	Ala	Pro	Thr	Ser	Arg	Trp	Gly	Thr	Pro	Asp	Gln
305					310					315					320
Phe	Arg	Ala	Leu	Val	Asp	Ala	Phe	His	Ala	Arg	Gly	Ile	Gly	Val	Ile
				325						330					335
Met	Asp	Trp	Val	Pro	Ala	His	Phe	Pro	Lys	Asp	Asp	Trp	Ala	Leu	Ala
			340						345						350
Arg	Phe	Asp	Gly	Glu	Ala	Leu	Tyr	Glu	His	Pro	Asp	Trp	Arg	Arg	Gly
		355						360					365		
Glu	Gln	Lys	Asp	Trp	Gly	Thr	Leu	Val	Phe	Asp	Phe	Gly	Arg	Asn	Glu
370						375						380			
Val	Arg	Asn	Phe	Leu	Val	Ala	Asn	Ala	Leu	Tyr	Trp	Ile	Glu	Glu	Phe
385					390						395				400
His	Ile	Asp	Gly	Leu	Arg	Val	Asp	Ala	Val	Ala	Ser	Met	Leu	Tyr	Leu
				405						410					415
Asp	Tyr	Ser	Arg	Glu	His	Gly	Glu	Trp	Glu	Pro	Asn	Ile	Tyr	Gly	Gly
			420						425					430	
Arg	Glu	Asn	Leu	Glu	Ala	Val	Gln	Phe	Leu	Gln	Glu	Met	Asn	Ala	Thr
		435						440						445	
Val	Leu	Arg	Leu	His	Pro	Gly	Ala	Leu	Thr	Ile	Ala	Glu	Glu	Ser	Thr
	450					455						460			
Ser	Trp	Pro	Gly	Val	Thr	Ala	Pro	Thr	Trp	Asp	Gly	Gly	Leu	Gly	Phe
465					470					475					480
Ser	Leu	Lys	Trp	Asn	Met	Gly	Trp	Met	His	Asp	Thr	Leu	Glu	Tyr	Phe
				485						490					495
Ser	Lys	Asn	Pro	Val	His	Arg	Ala	Phe	His	His	Ser	Glu	Leu	Thr	Phe
			500						505						510
Ser	Leu	Val	Tyr	Ala	Phe	Ser	Glu	Arg	Phe	Val	Leu	Pro	Ile	Ser	His
		515						520					525		
Asp	Glu	Val	Val	His	Gly	Lys	Gly	Ser	Leu	Trp	Asp	Arg	Met	Pro	Gly
	530					535							540		
Asp	Thr	Trp	Asn	Lys	Ala	Ala	Gly	Leu	Arg	Thr	Phe	Leu	Ala	Tyr	Met
545					550					555					560
Trp	Ser	His	Pro	Gly	Lys	Lys	Leu	Leu	Phe	Met	Gly	Gln	Glu	Phe	Gly
				565						570					575
Gln	Arg	Glu	Glu	Trp	Ala	Glu	Gly	Gln	Gly	Leu	Pro	Trp	Asp	Ile	Val
			580						585						590
Asp	Gly	Trp	Gln	Gly	Glu	Tyr	His	Glu	Ala	Ile	Arg	Thr	Leu	Thr	Arg
		595						600						605	
Ser	Leu	Asn	Gly	Val	Tyr	Ser	Asp	Ser	Pro	Ala	Leu	His	Thr	Gln	Asp
	610						615								620
Phe	Thr	Gly	Glu	Gly	Phe	Thr	Trp	Asn	Lys	Gly	Asp	Asp	Ala	Thr	Asn
625					630						635				640
Asn	Ile	Leu	Ala	Phe	Thr	Arg	Phe	Gly	Ser	Asp	Gly	Ser	Gln	Met	Leu
				645						650					655

-continued

Cys	Val	Phe	Asn	Leu	Ser	Gly	Thr	Ser	Gln	Pro	Glu	Tyr	Gln	Leu	Gly
			660					665					670		
Val	Ala	Ala	Gly	Gly	Glu	Trp	Lys	Leu	Val	Leu	Asn	Thr	Asp	Asp	Ala
			675				680					685			
Glu	Phe	Leu	Gly	Ala	Glu	Asn	Asp	Ile	Ala	Thr	Ser	Val	Gln	Ala	Ala
			690			695					700				
Ala	Thr	Pro	Arg	Asp	Asn	Phe	Ala	Tyr	Ser	Leu	Ser	Leu	His	Val	Pro
705					710					715					720
Ala	Met	Ser	Ala	Gln	Phe	Tyr	Ser	Leu	Gln	Lys					
				725					730						

What is claimed is:

1. A recombinant methanotrophic bacterium, comprising an exogenous nucleic acid that encodes a gluconeogenesis enzyme,

wherein the encoded gluconeogenesis enzyme is one or more of a pyruvate carboxylase, a phosphoenolpyruvate carboxykinase, an enolase, a phosphoglycerate mutase, a phosphoglycerate kinase, a glyceraldehyde-3-phosphate dehydrogenase, a Type A aldolase, a fructose 1,6-bisphosphatase, a phosphofructokinase, a phosphoglucose isomerase, a hexokinase, and a glucose-6-phosphate,

wherein the methanotrophic bacterium is selected from the group consisting of *Methylomonas*, *Methylobacter*, *Methylococcus*, *Methylosinus*, *Methylocystis*, *Methylo-microbium*, *Methanomonas*, and *Methylocella*, and

wherein the recombinant methanotrophic bacterium grown on a C₁ substrate feedstock is capable of producing glucose at a level that is greater than that produced by the parent methanotrophic bacterium.

2. The recombinant methanotrophic bacterium of claim 1, wherein the C₁ substrate feedstock comprises natural gas or methane.

3. The recombinant methanotrophic bacterium of claim 1, wherein the methanotrophic bacterium is an obligate methanotrophic bacterium.

4. The recombinant methanotrophic bacterium of claim 1, wherein the methanotrophic bacterium is a facultative methanotrophic bacterium.

5. The recombinant methanotrophic bacterium of claim 1, wherein the methanotrophic bacterium is selected from the group consisting of *Methylococcus capsulatus* Bath, *Methylomonas* sp. 16a, *Methylosinus trichosporium* OB3b, *Methylosinus sporium*, *Methylocystis parvus*, *Methylomonas methanica*, *Methylomonas albus*, *Methylobacter capsulatus* Y, *Methylomonas flagellata* AJ-3670, *Methylacidiphilum infernorum*, *Methylacidiphilum fumariolicum*, *Methylomicrobium alcaliphilum*, and *Methyloacida kamchatkensis*.

6. The recombinant methanotrophic bacterium of claim 1, wherein the exogenous nucleic acid encoding the gluconeogenesis enzyme is endogenous to an organism selected from the group consisting of a bacterium, a yeast, a fungi, and a plant.

7. The recombinant methanotrophic bacterium of claim 1, wherein the exogenous nucleic acid comprises an expression control sequence that is operably linked to the nucleic acid encoding the gluconeogenesis enzyme.

8. The recombinant methanotrophic bacterium of claim 1, wherein the exogenous nucleic acid encoding the gluconeogenesis enzyme is endogenous to *Escherichia coli* or *Corynebacterium glutamicum*.

9. The recombinant methanotrophic bacterium of claim 1, wherein the exogenous nucleic acid encoding the gluconeogenesis enzyme is endogenous to *Saccharomyces cerevisiae*.

10. The recombinant methanotrophic bacterium of claim 1, wherein the exogenous nucleic acid encoding the gluconeogenesis enzyme is codon optimized for the methanotrophic bacterium.

11. The recombinant methanotrophic bacterium of claim 1, wherein the carbohydrates of the recombinant methanotrophic bacterium exhibits a $\delta^{13}\text{C}$ that is less than -30% or is less than -40% .

12. The recombinant methanotrophic bacterium of claim 1, wherein the methanotrophic bacterium further comprises an exogenous polynucleotide encoding one or more glycogenesis enzymes selected from the group consisting of a glucose-1-phosphate adenylyltransferase, a glycogen synthase, and a 1,4-alpha-glucan-branching protein.

13. The recombinant methanotrophic bacterium of claim 12, wherein the further exogenous polynucleotide encoding one or more glycogenesis enzymes comprises an expression control sequence that is operably linked to the polynucleotide encoding the one or more glycogenesis enzymes, wherein the one or more glycogenesis enzymes are heterologous glycogenesis enzymes, native glycogenesis enzymes, or a combination thereof.

14. The recombinant methanotrophic bacterium of claim 12, wherein the exogenous nucleic acid encoding the one or more glycogenesis enzymes is codon optimized for the methanotrophic bacterium.

15. A biomass derived from whole and/or lysed cells of the recombinant methanotrophic bacterium of claim 1.

16. A carbohydrate composition, comprising carbohydrates extracted from a biomass derived from the methanotrophic bacterium of claim 1, wherein the composition exhibits a $\delta^{13}\text{C}$ that is less than -30% or is less than -40% .

17. An animal feed, comprising the recombinant methanotrophic bacterium of claim 1, a biomass derived from whole and/or lysed cells of the recombinant methanotrophic bacterium of claim 1, or a carbohydrate composition comprising carbohydrates extracted from a biomass derived from the recombinant methanotrophic bacterium of claim 1.

18. The animal feed of claim **17**, further comprising an additive selected from the group consisting of a plant-derived material, an animal-derived material, and a micro-organism-derived material.

19. The animal feed of claim **18**, wherein the additive is plant-derived material and the plant-derived material is selected from the group consisting of corn, soybean meal, pea protein, or a combination thereof.

20. The animal feed of claim **18**, wherein the additive is an animal-derived material and the animal-derived material is fish meal.

21. A method of producing a desired carbohydrate, the method comprising culturing the recombinant methanotrophic bacterium of claim **1** in the presence of a C₁ substrate feedstock comprising methane under conditions sufficient to produce the glucose.

22. The method of claim **21**, wherein the natural gas-derived carbon feedstock is natural gas.

23. The method of claim **21**, wherein the methanotrophic bacterium is selected from the group consisting of *Methylococcus capsulatus* Bath, *Methylomonas* sp. 16a, *Methylosinus trichosporium* OB3b, *Methylosinus sporium*, *Methylocystis parvus*, *Methylomonas methanica*, *Methylomonas albus*, *Methylobacter capsulatus* Y, *Methylomonas flagellata* AJ-3670, *Methylacidiphilum infernorum*, *Methylacidiphilum fumariolicum*, *Methylomicrobium alcaliphilum*, and *Methyloacida kamchatkensis*.

* * * * *