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(54) Title: CHIMERIC ABC TRANSPORTERS AND SCREENING METHODS

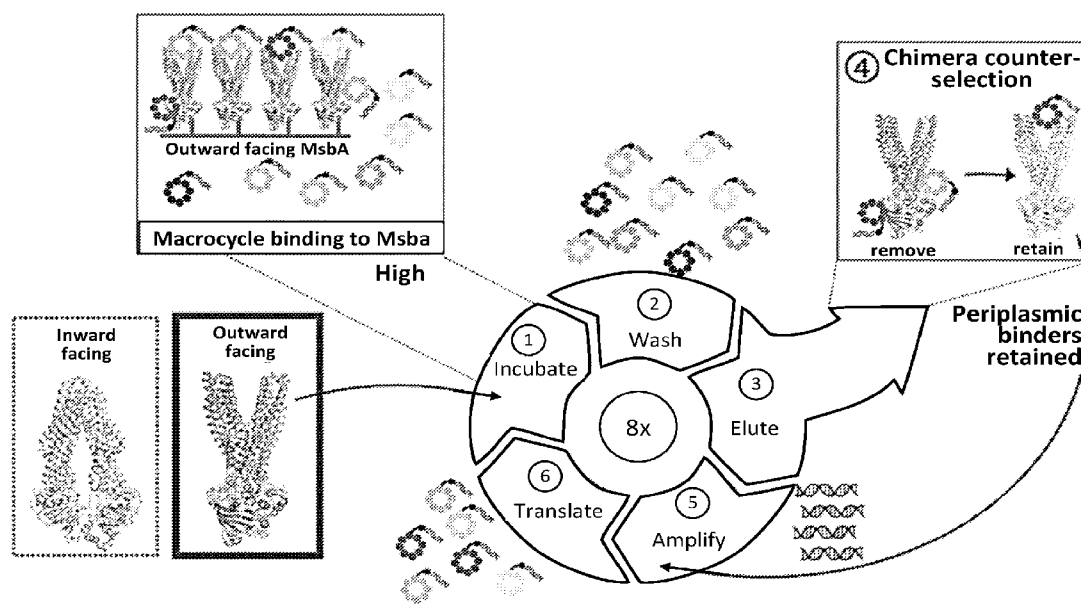


Fig. 3A

(57) Abstract: The present disclosure relates to chimeric ABC transporter proteins and methods of screening for molecules that bind to the periplasmic, extracellular, and/or luminal face of an ABC transporter protein using the chimeric ABC transporters. For example, in some embodiments, screening methods involve providing a chimeric ABC transporter in which one or more regions of the periplasmic, extracellular, and/or luminal face of the ABC transporter are substituted with one or more equivalent regions of the periplasmic, extracellular, and/or luminal face of a different ABC transporter and selecting for molecules that bind to the ABC transporter but do not bind to the chimeric ABC transporter. The disclosure also relates to molecules that bind to the periplasmic, extracellular, and/or luminal face of an ABC transporter protein, for example, identified in such screens.



(74) **Agent: DOHERTY, Elizabeth A.** et al.; 125 Cambridge Park Drive, Suite 301, Cambridge, Massachusetts 02140 (US).

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A. CLASSIFICATION OF SUBJECT MATTER				
INV. G01N33/68	A61K39/00	C07K14/00		
C12Q1/18	C07K7/00	C12N15/00		
C07K16/00	C12N9/00			
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According to International Patent Classification (IPC) or to both national classification and IPC				
B. FIELDS SEARCHED				
Minimum documentation searched (classification system followed by classification symbols) G01N C07K A61K C12N C12Q				
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched				
Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) EPO-Internal, BIOSIS, EMBASE, WPI Data				
C. DOCUMENTS CONSIDERED TO BE RELEVANT				
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.		
X	<p>TEEM J L ET AL: "Identification of revertants for the cystic fibrosis @DF508 mutation using STE6-CFTR chimeras in yeast", CELL, ELSEVIER, AMSTERDAM NL, vol. 73, no. 2, 23 April 1993 (1993-04-23) , pages 335-346, XP024245690, ISSN: 0092-8674, DOI: 10.1016/0092-8674(93)90233-G [retrieved on 1993-04-23] whole document, in particular fig. 1; p. 336, col. 1, par. 1-3; p. 344, col. 1, par. 1, 2; col. 2, par. 1, 2</p> <p style="text-align: center;">----- -/--</p>	65, 67-69		
<table style="width: 100%; border: none;"> <tr> <td style="width: 50%; border: none;"><input checked="" type="checkbox"/> Further documents are listed in the continuation of Box C.</td> <td style="width: 50%; border: none;"><input checked="" type="checkbox"/> See patent family annex.</td> </tr> </table>			<input checked="" type="checkbox"/> Further documents are listed in the continuation of Box C.	<input checked="" type="checkbox"/> See patent family annex.
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Date of the actual completion of the international search	Date of mailing of the international search report			
16 December 2022	02/01/2023			
Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Fax: (+31-70) 340-3016	Authorized officer Chrétien, Eva Maria			

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US2022/026319

Box No. I Nucleotide and/or amino acid sequence(s) (Continuation of item 1.c of the first sheet)

1. With regard to any nucleotide and/or amino acid sequence disclosed in the international application, the international search was carried out on the basis of a sequence listing:
 - a. forming part of the international application as filed:
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 - on paper or in the form of an image file.
 - b. furnished together with the international application under PCT Rule 13ter.1(a) for the purposes of international search only in the form of an Annex C/ST.25 text file.
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INTERNATIONAL SEARCH REPORT

International application No
PCT/US2022/026319

C(Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	<p>ALEXANDER MARY KATE ET AL: "Disrupting Gram-Negative Bacterial Outer Membrane Biosynthesis through Inhibition of the Lipopolysaccharide Transporter MsbA", ANTIMICROBIAL AGENTS AND CHEMOTHERAPY, vol. 62, no. 11, 13 August 2018 (2018-08-13), XP055960408, US ISSN: 0066-4804, DOI: 10.1128/AAC.01142-18 Retrieved from the Internet: URL:https://journals.asm.org/doi/pdf/10.1128/AAC.01142-18> whole document, in particular p. 6-7, bridging par.; fig. 3; table 4</p>	65, 67-78, 81,83
X	<p>GUO XIAOXIAN ET AL: "A Mutation in Intracellular Loop 4 Affects the Drug-Efflux Activity of the Yeast Multidrug Resistance ABC Transporter Pdr5p", PLOS ONE, vol. 7, no. 1, 6 January 2012 (2012-01-06), page e29520, XP093008668, DOI: 10.1371/journal.pone.0029520 Retrieved from the Internet: URL:https://journals.plos.org/plosone/article/file?id=10.1371/journal.pone.0029520&type=printable> whole document, in particular p. 1, col. 2, bridging par. - p. 3, col. 1, par. 3; fig. 1, 2</p>	65-69
X	<p>SACHS G ET AL: "Urea Transport in Bacteria: Acid Acclimation by Gastric Helicobacter spp", JOURNAL OF MEMBRANE BIOLOGY, SPRINGER-VERLAG, NE, vol. 212, no. 2, 30 January 2007 (2007-01-30), pages 71-82, XP019489232, ISSN: 1432-1424 whole document, in particular p. 78, col. 2, par. 2</p>	65, 67-70,77
A	<p>WO 2005/064021 A2 (GLAXOSMITHKLINE BIOLOG SA [BE]; UNIV UTRECHT [NL] ET AL.) 14 July 2005 (2005-07-14) the whole document</p>	65-83

INTERNATIONAL SEARCH REPORT

International application No PCT/US2022/026319
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C(Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	<p>VAKKASOGLU AHMET S. ET AL: "D-helix influences dimerization of the ATP-binding cassette (ABC) transporter associated with antigen processing 1 (TAP1) nucleotide-binding domain", PLOS ONE, vol. 12, no. 5, 23 May 2017 (2017-05-23), page e0178238, XP093008700, DOI: 10.1371/journal.pone.0178238 Retrieved from the Internet: URL:https://journals.plos.org/plosone/article/file?id=10.1371/journal.pone.0178238&type=printable> the whole document</p>	65-83
A	<p align="center">-----</p> <p>RAETZ C R H ET AL: "LIPOPOLYSACCHARIDE ENDOTOXINS", ANNUAL REVIEW OF BIOCHEMISTRY, PALTO ALTO, CA, US, vol. 71, 1 January 2002 (2002-01-01), pages 635-700, XP001155986, ISSN: 0066-4154, DOI: 10.1146/ANNUREV.BIOCHEM.71.110601.135414 cited in the application the whole document</p> <p align="center">-----</p>	65-83

INTERNATIONAL SEARCH REPORT

Information on patent family members

International application No

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Patent document cited in search report	Publication date	Patent family member(s)	Publication date	
WO 2005064021	A2	14-07-2005	CA 2550927 A1	14-07-2005
			EP 1706481 A2	04-10-2006
			JP 5173194 B2	27-03-2013
			JP 2007515180 A	14-06-2007
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