



- (51) International Patent Classification:  
*G06F 19/22* (2011.01) *C12Q 1/68* (2006.01)
- (21) International Application Number:  
PCT/US2013/043123
- (22) International Filing Date:  
29 May 2013 (29.05.2013)
- (25) Filing Language: English
- (26) Publication Language: English
- (30) Priority Data:  
61/652,784 29 May 2012 (29.05.2012) US
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- (74) Agents: ELRIFI, Ivor, R. et al.; Mintz Levin Cohn Ferris Glovsky and Popeo, P.C., Chrysler Center, 666 Third Avenue, New York, NY 10017 (US).
- (81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM,

AO, AT, AU, AZ, BA, BB, BG, BH, BN, BR, BW, BY, BZ, CA, CH, CL, CN, CO, CR, CU, CZ, DE, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PA, PE, PG, PH, PL, PT, QA, RO, RS, RU, RW, SC, SD, SE, SG, SK, SL, SM, ST, SV, SY, TH, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW.

(84) Designated States (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, GH, GM, KE, LR, LS, MW, MZ, NA, RW, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, RU, TJ, TM), European (AL, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HR, HU, IE, IS, IT, LT, LU, LV, MC, MK, MT, NL, NO, PL, PT, RO, RS, SE, SI, SK, SM, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, KM, ML, MR, NE, SN, TD, TG).

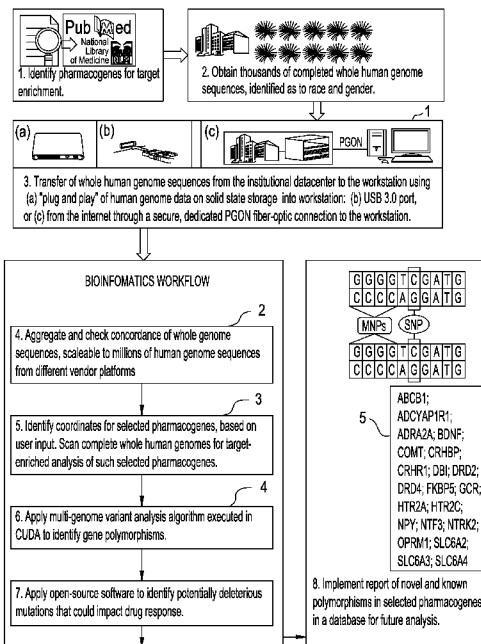
Published:

— with international search report (Art. 21(3))

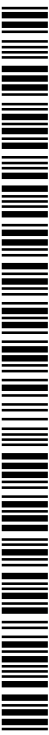
(88) Date of publication of the international search report:  
17 July 2014

(54) Title: NOVEL PHARMACOGENE SINGLE NUCLEOTIDE POLYMORPHISMS AND METHODS OF DETECTING SAME

FIG. 1



(57) Abstract: The present invention provides pharmacogene polymorphisms and their use in predicting therapeutic effectiveness. The present invention also provides methods comprising targeted analysis of selected pharmacogenes in thousands of compiled whole human genome sequences for identifying polymorphic sequences associated with drug response are described. The methods also provide confirmation and validation of these pharmacogene polymorphisms, based on concordance between different sequencing technologies, and statistical error-checking. Imputation of the deleterious consequences of novel variants is predicted by bioinformatics analysis.



# INTERNATIONAL SEARCH REPORT

International application No PCT/US2013/043123
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**A. CLASSIFICATION OF SUBJECT MATTER**  
 INV. G06F19/22 C12Q1/68  
 ADD.

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)  
 G06F 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, FSTA, WPI Data

<b>C. DOCUMENTS CONSIDERED TO BE RELEVANT</b>		
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	SHI MICHAEL M: "Enabling large-scale pharmacogenetic studies by high-throughput mutation detection and genotyping technologies", CLINICAL CHEMISTRY, AMERICAN ASSOCIATION FOR CLINICAL CHEMISTRY, WASHINGTON, DC, vol. 47, no. 2, 1 February 2000 (2000-02-01), pages 164-172, XP002197957, ISSN: 0009-9147 abstract page 165, right-hand column, paragraph 1st page 166, paragraph bridging - page 167 ----- -/--	1-4

<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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* Special categories of cited documents :  "A" document defining the general state of the art which is not considered to be of particular relevance "E" earlier application or patent but published on or after the international filing date "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed	"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention  "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art  "&" document member of the same patent family
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Date of the actual completion of the international search  <p style="text-align: center; font-size: 1.2em;">27 May 2014</p>	Date of mailing of the international search report  <p style="text-align: center; font-size: 1.2em;">04/06/2014</p>
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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  <p style="text-align: center; font-size: 1.2em;">Ripaud, Leslie</p>
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**INTERNATIONAL SEARCH REPORT**

International application No PCT/US2013/043123
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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.
Y	<p>MIAN LU ET AL: "G SNP: A DNA Single-Nucleotide Polymorphism Detection System with GPU Acceleration", PARALLEL PROCESSING (ICPP), 2011 INTERNATIONAL CONFERENCE ON, IEEE, 13 September 2011 (2011-09-13), pages 592-601, XP032461059, DOI: 10.1109/ICPP.2011.51 ISBN: 978-1-4577-1336-1 abstract page 598, left-hand column page 601, left-hand column, line 1st full</p> <p align="center">-----</p>	1-4
X	<p>CONSOLI GIORGIO ET AL: "ABC B1 polymorphisms are associated with clozapine plasma levels in psychotic patients", PHARMACOGENOMICS, FUTURE MEDICINE LTD, UK, vol. 10, no. 8, 1 August 2009 (2009-08-01), pages 1267-1276, XP009175493, ISSN: 1744-8042, DOI: 10.2217/PGS.09.51 abstract page 1267, left-hand column, paragraph 1st</p> <p align="center">-----</p>	5,6
A	<p>MARTH G T ET AL: "GENERAL APPROACH TO SINGLE-NUCLEOTIDE POLYMORPHISM DISCOVERY", NATURE GENETICS, NATURE PUBLISHING GROUP, NEW YORK, US, vol. 23, no. 4, 1 December 1999 (1999-12-01), pages 452-456, XP000920898, ISSN: 1061-4036, DOI: 10.1038/70570 abstract</p> <p align="center">-----</p>	1-4
A	<p>R. R. FREIMUTH ET AL: "Polymorphism discovery in 51 chemotherapy pathway genes", HUMAN MOLECULAR GENETICS, vol. 14, no. 23, 3 November 2005 (2005-11-03), pages 3595-3603, XP055096623, ISSN: 0964-6906, DOI: 10.1093/hmg/ddi387 abstract</p> <p align="center">-----</p> <p align="center">-/--</p>	1-4

## INTERNATIONAL SEARCH REPORT

International application No  
PCT/US2013/043123

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>LINDA L PELLEYMOUNTER ET AL: "A novel application of pattern recognition for accurate SNP and indel discovery from high-throughput data: Targeted resequencing of the glucocorticoid receptor co-chaperone FKBP5 in a Caucasian population", MOLECULAR GENETICS AND METABOLISM, ACADEMIC PRESS, SAN DIEGO, CA, US, vol. 104, no. 4, 18 August 2011 (2011-08-18), pages 457-469, XP028117208, ISSN: 1096-7192, DOI: 10.1016/J.YMGME.2011.08.019 [retrieved on 2011-08-24] abstract</p> <p style="text-align: center;">-----</p>	1-4
A	<p>K IRIZARRY ET AL: "Single nucleotide polymorphism identification in candidate gene systems of obesity", THE PHARMACOGENOMICS JOURNAL, vol. 1, no. 3, 1 January 2001 (2001-01-01), pages 193-203, XP055096633, ISSN: 1470-269X, DOI: 10.1038/sj.tpj.6500042 abstract page 194, right-hand column</p> <p style="text-align: center;">-----</p>	1-4
A	<p>RASMUS NIELSEN ET AL: "Genotype and SNP calling from next-generation sequencing data", NATURE REVIEWS GENETICS, vol. 12, no. 6, 1 June 2011 (2011-06-01), pages 443-451, XP055046801, ISSN: 1471-0056, DOI: 10.1038/nrg2986 the whole document</p> <p style="text-align: center;">-----</p>	1-4
A	<p>HOMANN OLIVER R ET AL: "MochiView: versatile software for genome browsing and DNA motif analysis", BMC BIOLOGY, BIOMED CENTRAL, LONDON, GB, GB, vol. 8, no. 1, 21 April 2010 (2010-04-21), page 49, XP021079396, ISSN: 1741-7007, DOI: 10.1186/1741-7007-8-49 cited in the application the whole document</p> <p style="text-align: center;">-----</p> <p style="text-align: center;">-/--</p>	1-4

## INTERNATIONAL SEARCH REPORT

International application No  
PCT/US2013/043123

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>Jan Van Oeveren ET AL: "Mining SNPs from DNA Sequence Data; Computational Approaches to SNP Discovery and Analysis", Single Nucleotide Polymorphisms, Methods in Molecular Biology 578, 1 January 2009 (2009-01-01), pages 73-91, XP55071076, DOI: 10.1007/978-1-60327-411-1_4,a Retrieved from the Internet: URL:http://download.bioon.com.cn/view/upload/month_1004/20100419_ee17b59a19517c3eb17cIBjUuh9eoYMF.attach.pdf [retrieved on 2013-07-12] the whole document</p>	1-4
A	<p>US 2003/211504 A1 (FECHTEL KIM [US] ET AL) 13 November 2003 (2003-11-13) paragraphs [0009], [0010], [0032]</p>	1-4
X	<p>Takao Isogai: "EM_EST:DA227565", 18 October 2005 (2005-10-18), XP055118688, Retrieved from the Internet: URL:http://ibis/exam/dbfetch.jsp?id=EM_EST:DA227565 [retrieved on 2014-05-19] the whole document</p>	7-10
X	<p>KIMURA KOUICHI ET AL: "Diversification of transcriptional modulation: large-scale identification and characterization of putative alternative promoters of human genes", GENOME RESEARCH, COLD SPRING HARBOR LABORATORY PRESS, WOODBURY, NY, US, vol. 16, no. 1, 1 January 2006 (2006-01-01), pages 55-65, XP002446089, ISSN: 1088-9051, DOI: 10.1101/GR.4039406 page 56, right-hand column, paragraph 2nd page 63, right-hand column, paragraph 1st</p>	7-10
X	<p>Sung-Jong Hong: "EM_EST:AT009657", 13 September 2006 (2006-09-13), XP055118695, Retrieved from the Internet: URL:http://ibis/exam/dbfetch.jsp?id=EM_EST:AT009657 [retrieved on 2014-05-19] the whole document</p>	7-10
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## INTERNATIONAL SEARCH REPORT

International application No

PCT/US2013/043123

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>PYO YUN CHO ET AL: "Expressed sequence tag analysis of adult Clonorchis sinensis, the Chinese liver fluke",            PARASITOLOGY RESEARCH ; FOUNDED AS            ZEITSCHRIFT FÜR PARASITENKUNDE, SPRINGER,            BERLIN, DE,            vol. 99, no. 5, 17 May 2006 (2006-05-17),            pages 602-608, XP019427994,            ISSN: 1432-1955, DOI:            10.1007/S00436-006-0204-1            page 603, paragraph bridging - page 604            -----</p>	7-10
X	<p>Carninci ET AL: "EM_EST:BY315426",            17 December 2002 (2002-12-17),            XP055118808,            Retrieved from the Internet:            URL:http://ibis/exam/dbfetch.jsp?id=EM_EST            :BY315426            [retrieved on 2014-05-20]            the whole document            -----</p>	7-10
X	<p>Carninci ET AL: "EM_EST:BY306447",            17 December 2002 (2002-12-17),            XP055118810,            Retrieved from the Internet:            URL:http://ibis/exam/dbfetch.jsp?id=EM_EST            :BY306447            [retrieved on 2014-05-20]            the whole document            -----</p>	7-10
X	<p>CARNINCI P ET AL: "NORMALIZATION AND            SUBTRACTION OF CAP-TRAPPER-SELECTED CDNAS            TO PREPARE FULL-LENGTH CDNA LIBRARIES FOR            RAPID DISCOVERY OF NEW GENES",            GENOME RESEARCH, COLD SPRING HARBOR            LABORATORY PRESS, WOODBURY, NY, US,            vol. 10, no. 10,            1 January 2000 (2000-01-01), pages            1617-1630, XP002944079,            ISSN: 1088-9051, DOI: 10.1101/GR.145100            page 1628, right-hand column, last            paragraph            -----            -/--</p>	7-10

## INTERNATIONAL SEARCH REPORT

International application No  
PCT/US2013/043123

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>MINGQING XU ET AL: "Genetic variants in the BDNF gene and therapeutic response to risperidone in schizophrenia patients: a pharmacogenetic study", EUROPEAN JOURNAL OF HUMAN GENETICS, vol. 18, no. 6, 20 January 2010 (2010-01-20), pages 707-712, XP055118741, ISSN: 1018-4813, DOI: 10.1038/ejhg.2009.238 abstract page 707, left-hand column, paragraph 1st -----</p>	5,6
X	<p>Pj Blackshear: "EM_EST:AW638725", 5 April 2000 (2000-04-05), XP055119062, Retrieved from the Internet: URL:http://ibis/exam/dbfetch.jsp?id=EM_EST:AW638725 [retrieved on 2014-05-20] the whole document -----</p>	7-10
X	<p>Dk Fisher ET AL: "GSN:ARP47336", 21 August 2008 (2008-08-21), XP055119064, Retrieved from the Internet: URL:http://ibis/exam/dbfetch.jsp?id=GSN:ARP47336 [retrieved on 2014-05-20] the whole document -----</p>	7-10
X	<p>Jc Venter ET AL: "GSN:AFI74404", 2 November 2004 (2004-11-02), XP055119136, Retrieved from the Internet: URL:http://ibis/exam/dbfetch.jsp?id=GSN:AFI74404 [retrieved on 2014-05-21] the whole document -----</p>	7
X	<p>ELISABETH B. BINDER: "Association of Polymorphisms in Genes Regulating the Corticotropin-Releasing Factor System With Antidepressant Treatment Response", ARCHIVES OF GENERAL PSYCHIATRY, vol. 67, no. 4, 1 April 2010 (2010-04-01), page 369, XP055075005, ISSN: 0003-990X, DOI: 10.1001/archgenpsychiatry.2010.18 abstract ----- -/--</p>	5,6

## INTERNATIONAL SEARCH REPORT

International application No  
PCT/US2013/043123

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>R. Strausberg: "EM_EST:AI418253",  <sup>3</sup>            12 March 1999 (1999-03-12), XP055119169,            Retrieved from the Internet:            URL:http://ibis/exam/dbfetch.jsp?id=EM_EST:            AI418253            [retrieved on 2014-05-21]            the whole document</p> <p style="text-align: center;">-----</p>	7-10
X	<p>Yc Kim ET AL: "EM_EST:DR978925",  <sup>3</sup>            5 December 2005 (2005-12-05), XP055119173,            Retrieved from the Internet:            URL:http://ibis/exam/dbfetch.jsp?id=EM_EST:            DR978925            [retrieved on 2014-05-21]            the whole document</p> <p style="text-align: center;">-----</p>	7-10
X	<p>M Suwa ET AL: "GSN:ADC86254",  <sup>3</sup>            11 June 2007 (2007-06-11), XP055119179,            Retrieved from the Internet:            URL:http://ibis/exam/dbfetch.jsp?id=GSN:AD            C86254            [retrieved on 2014-05-21]            the whole document</p> <p style="text-align: center;">-----</p>	7
X	<p>LICINIO J ET AL: "Association of a            corticotropin-releasing hormone receptor 1            haplotype and antidepressant treatment            response in Mexican-Americans",            MOLECULAR PSYCHIATRY, BASINGSTOKE, GB,            vol. 9, no. 12,            1 January 2004 (2004-01-01), pages            1075-1082, XP002540580,            ISSN: 1359-4184, DOI:            10.1038/SJ.MP.4001587            [retrieved on 2004-09-14]            abstract</p> <p style="text-align: center;">-----</p>	5,6
X	<p>J Kalicki: "EM_STD:AC004822",  <sup>3</sup>            15 June 1998 (1998-06-15), XP055119349,            Retrieved from the Internet:            URL:http://ibis/exam/dbfetch.jsp?id=EM_STD:            AC004822            [retrieved on 2014-05-21]            the whole document</p> <p style="text-align: center;">-----</p> <p style="text-align: center;">-/--</p>	7,9,10

## INTERNATIONAL SEARCH REPORT

International application No  
PCT/US2013/043123

C(Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	Am Ozyildirim ET AL: "EM_EST:CK430028", 7 January 2004 (2004-01-07), XP055119350, Retrieved from the Internet: URL:http://ibis/exam/dbfetch.jsp?id=EM_EST: CK430028 [retrieved on 2014-05-21] the whole document	7-10
X	Gg Mahairas ET AL: "EM_GSS:AQ134147", 8 September 1998 (1998-09-08), XP055119352, Retrieved from the Internet: URL:http://ibis/exam/dbfetch.jsp?id=EM_GSS: AQ134147 [retrieved on 2014-05-21] the whole document	7,9,10
X	B R GODLEWSKA ET AL: "0lanzapine-induced weight gain is associated with the -759C/T and -697G/C polymorphisms of the HTR2C gene", THE PHARMACOGENOMICS JOURNAL, vol. 9, no. 4, 12 May 2009 (2009-05-12), pages 234-241, XP055119354, ISSN: 1470-269X, DOI: 10.1038/tpj.2009.18 abstract page 234, paragraph 1st	5,6
X	Ll Moroz ET AL: "EM_EST:GR416073", 13 May 2010 (2010-05-13), XP055119409, Retrieved from the Internet: URL:http://ibis/exam/dbfetch.jsp?id=EM_EST: GR416073 [retrieved on 2014-05-22] the whole document	7-10
X	H Beasley: "EM_STD:AL596132", 18 July 2001 (2001-07-18), XP055119413, Retrieved from the Internet: URL:http://ibis/exam/dbfetch.jsp?id=EM_STD: AL596132 [retrieved on 2014-05-22] the whole document	7,9,10
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## INTERNATIONAL SEARCH REPORT

International application No  
PCT/US2013/043123

C(Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	J Aerssens ET AL: "GSN:AE61704", <sup>3</sup> 9 February 2006 (2006-02-09), XP055119415, Retrieved from the Internet: URL:http://ibis/exam/dbfetch.jsp?id=GSN:AE E61704 [retrieved on 2014-05-22] the whole document -----	7
X	C DONG ET AL: "Sequence variations of ABCB1, SLC6A2, SLC6A3, SLC6A4, CREB1, CRHR1 and NTRK2: association with major depression and antidepressant response in Mexican-Americans", MOLECULAR PSYCHIATRY, vol. 14, no. 12, 20 December 2009 (2009-12-20), pages 1105-1118, XP055075189, ISSN: 1359-4184, DOI: 10.1038/mp.2009.92 abstract page 1110, left-hand column, paragraph bridging - right-hand column -----	5,6
X	P Porzgen: "EM_STD:X91126", <sup>3</sup> 21 December 1995 (1995-12-21), XP055119475, Retrieved from the Internet: URL:http://ibis/exam/dbfetch.jsp?id=EM_STD :X91126 [retrieved on 2014-05-22] the whole document -----	7-10
A	Anonymous: "Reference SNP (refSNP) Cluster Report: rs201736463", dbSNP <sup>2</sup> 2 March 2012 (2012-03-02), XP055119958, Retrieved from the Internet: URL:http://www.ncbi.nlm.nih.gov/projects/S NP/snp_ref.cgi?rs=201736463 [retrieved on 2014-05-26] the whole document -----	1-10
A	Anonymous: "Reference SNP (refSNP) Cluster Report: rs113565135", dbSNP <sup>3</sup> 16 February 2010 (2010-02-16), XP055119960, Retrieved from the Internet: URL:http://www.ncbi.nlm.nih.gov/projects/S NP/snp_ref.cgi?rs=113565135 [retrieved on 2014-05-26] the whole document -----	1-10
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## INTERNATIONAL SEARCH REPORT

International application No

PCT/US2013/043123

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>Anonymous: "Reference SNP (refSNP) Cluster Report: rs74657068", dbSNP</p> <p>22 April 2010 (2010-04-22), XP055119961, Retrieved from the Internet: URL:http://www.ncbi.nlm.nih.gov/projects/SNP/snp_ref.cgi?rs=74657068 [retrieved on 2014-05-26] the whole document</p> <p>-----</p>	5-10
A,P	<p>Anonymous: "Reference SNP (refSNP) Cluster Report: rs370427318", dbSNP</p> <p>22 November 2012 (2012-11-22), XP055119962, Retrieved from the Internet: URL:http://www.ncbi.nlm.nih.gov/projects/SNP/snp_ref.cgi?rs=370427318 [retrieved on 2014-05-26] the whole document</p> <p>-----</p>	1-10
A	<p>Anonymous: "Reference SNP (refSNP) Cluster Report: rs146005473", dbSNP</p> <p>24 March 2011 (2011-03-24), XP055119964, Retrieved from the Internet: URL:http://www.ncbi.nlm.nih.gov/projects/SNP/snp_ref.cgi?rs=146005473 [retrieved on 2014-05-26] the whole document</p> <p>-----</p>	1-10
A	<p>Anonymous: "Reference SNP (refSNP) Cluster Report: rs187366360", dbSNP</p> <p>20 July 2011 (2011-07-20), XP055119966, Retrieved from the Internet: URL:http://www.ncbi.nlm.nih.gov/projects/SNP/snp_ref.cgi?rs=187366360 [retrieved on 2014-05-26] the whole document</p> <p>-----</p>	1-10
A	<p>Anonymous: "Reference SNP (refSNP) Cluster Report: rs200666890", dbSNP</p> <p>10 February 2012 (2012-02-10), XP055119968, Retrieved from the Internet: URL:http://www.ncbi.nlm.nih.gov/projects/SNP/snp_ref.cgi?rs=200666890 [retrieved on 2014-05-26] the whole document</p> <p>-----</p>	1-10

# INTERNATIONAL SEARCH REPORT

International application No.  
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## Box No. II Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)

This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1.  Claims Nos.:  
because they relate to subject matter not required to be searched by this Authority, namely:
  
2.  Claims Nos.:  
because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:
  
3.  Claims Nos.:  
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

## Box No. III Observations where unity of invention is lacking (Continuation of item 3 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

see additional sheet

1.  As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
  
2.  As all searchable claims could be searched without effort justifying an additional fees, this Authority did not invite payment of additional fees.
  
3.  As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:  
  
1-4(completely); 5-10(partially)
  
4.  No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

### Remark on Protest

- The additional search fees were accompanied by the applicant's protest and, where applicable, the payment of a protest fee.
- The additional search fees were accompanied by the applicant's protest but the applicable protest fee was not paid within the time limit specified in the invitation.
- No protest accompanied the payment of additional search fees.

**FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210**

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. claims: 1-4

concerns a method for interrogating thousands of aggregated whole human genome sequences comprising using a targeted analysis of one or more selected pharmacogenes and determining polymorphic sequences that may associate with a drug response.

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2. claims: 5-10(partially)

concerns a method for determining likelihood of an adverse or modified response to an anti-depressant or psychiatric drug comprising detecting the presence of a polymorphism, an isolated nucleic acid comprising said polymorphism, and a vector and a cell comprising said isolated nucleic acid, wherein said polymorphism is SEQ ID N° 1.

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3-119. claims: 5-10(partially)

idem invention 2, wherein inventions 3-119 are defined by the polymorphisms of SEQ ID N° 2-118, respectively.

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# INTERNATIONAL SEARCH REPORT

Information on patent family members

International application No

PCT/US2013/043123

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
US 2003211504	A1	NONE	
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