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(54) Title: POLYMORPHISMS IN THE HUMAN cMOAT GENE AND USES THEREOF

(57) Abstract: This invention relates to polymorphisms in the human cMOAT gene and corresponding novel allelic polypeptides encoded thereby. The invention also relates to methods and materials for analysing allelic variation in the cMOAT gene, and to the use of cMOAT polymorphism in treatment of diseases with cMOAT transportable drugs.

INTERNATIONAL SEARCH REPORT

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A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C12Q1/68 C07K16/18 C07K14/705		
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) IPC 7 C12Q C07K		
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 practical, search terms used) EPO-Internal, BIOSIS, MEDLINE, EMBL, PAJ, WPI Data, EMBASE, CHEM ABS Data		
C. DOCUMENTS CONSIDERED TO BE RELEVANT		
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WADA M ET AL: "Mutations in the canalicular multispecific organic anion transporter (cMOAT) gene, a novel ABC transporter, in patients with hyperbilirubinemia II/Dubin-Johnson syndrome." HUMAN MOLECULAR GENETICS. ENGLAND FEB 1998, vol. 7, no. 2, February 1998 (1998-02), pages 203-207, XP002249633 ISSN: 0964-6906	1,4,7
Y	the whole document --- -/--	2,5,6
<input checked="" type="checkbox"/> Further documents are listed in the continuation of box C. <input type="checkbox"/> Patent family members are listed in annex.		
° 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 document 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		
Date of the actual completion of the international search 31 July 2003		Date of mailing of the international search report 12/08/2003
Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016		Authorized officer Armando1a, E

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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.
X	TOH S ET AL: "Genomic structure of the canalicular multispecific organic anion-transporter gene (MRP2/cMOAT) and mutations in the ATP-binding-cassette region in Dubin-Johnson syndrome." AMERICAN JOURNAL OF HUMAN GENETICS. UNITED STATES MAR 1999, vol. 64, no. 3, March 1999 (1999-03), pages 739-746, XP002249635 ISSN: 0002-9297	1,4,7
Y	the whole document	2,5,6
X	KERB R ET AL: "ABC drug transporters: hereditary polymorphisms and pharmacological impact in MDR1, MRP1 and MRP2." PHARMACOGENOMICS. ENGLAND FEB 2001, vol. 2, no. 1, February 2001 (2001-02), pages 51-64, XP009014717 ISSN: 1462-2416	2,7
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X	DATABASE EMBL 'Online! 7 November 1996 (1996-11-07) TANIGUCHI K. ET AL.: "Canalicular multispecific organic anion transporter 'Homo sapiens!'" Database accession no. AAB39892 XP002249640 Sequence contains Val1188 the whole document -& TANIGUCHI KEN ET AL: "A human canalicular multispecific organic anion transporter (cMOAT) gene is overexpressed in cisplatin-resistant human cancer cell lines with decreased drug accumulation." CANCER RESEARCH, vol. 56, no. 18, 1996, pages 4124-4129, XP000607845 ISSN: 0008-5472	9
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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.
X	<p>TSUJII HIROYUKI ET AL: "Exon-intron organization of the human multidrug-resistance protein 2 (MRP2) gene mutated in Dubin-Johnson syndrome." GASTROENTEROLOGY, vol. 117, no. 3, 1999, pages 653-660, XP009014900 ISSN: 0016-5085 the whole document -& DATABASE EMBL 'Online! 11 June 2003 (2003-06-11) KEPPLER D.: "Multidrug resistance protein 2 (MRP2) (Homo sapiens)" Database accession no. CAB45309 XP002249641 Sequence contains Val1188 ---</p>	9
X	<p>BUECHLER M ET AL: "CDNA CLONING OF THE HEPATOCYTE CANALICULAR ISOFORM OF THE MULTIDRUG RESISTANCE PROTEIN, CMRP, REVEALS A NOVEL CONJUGATE EXPORT PUMP DEFICIENT IN HYPERBILIRUBINEMIC MUTANT RATS" JOURNAL OF BIOLOGICAL CHEMISTRY, AMERICAN SOCIETY OF BIOLOGICAL CHEMISTS, BALTIMORE, MD, US, vol. 271, no. 25, 21 June 1996 (1996-06-21), pages 15091-15098, XP000606392 ISSN: 0021-9258 -& DATABASE EMBL 'Online! 25 May 2001 (2001-05-25) KEPPLER D.: "Canalicular multidrug resistance protein (Homo sapiens)" Database accession no. CAA65259 XP002249642 Sequence contains Val1188 ---</p>	9
Y	<p>SCHEFFER GEORGE L ET AL: "Specific detection of multidrug resistance proteins MRP1, MRP2, MRP3, MRP5, and MDR3 P-glycoprotein with a panel of monoclonal antibodies." CANCER RESEARCH, vol. 60, no. 18, 15 September 2000 (2000-09-15), pages 5269-5277, XP002249637 ISSN: 0008-5472 table 1 --- -/--</p>	10

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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>PAULUSMA C C ET AL: "CONGENITAL JAUNIDICE IN RATS WITH A MUTATION IN A MULTIDRUG RESISTANCE-ASSOCIATED PROTEIN GENE" SCIENCE, AMERICAN ASSOCIATION FOR THE ADVANCEMENT OF SCIENCE,, US, vol. 271, 23 February 1996 (1996-02-23), pages 1126-1128, XP000606395 ISSN: 0036-8075 the whole document -----</p>	10