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A61K 31/4015 (2006.01) A61P 1/16 (2006.01)
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[Continued on next page]

(54) Title: COMPOSITIONS AND METHODS FOR REDUCING THE RISK OF AGENT-INDUCED LIVER TOXICITY

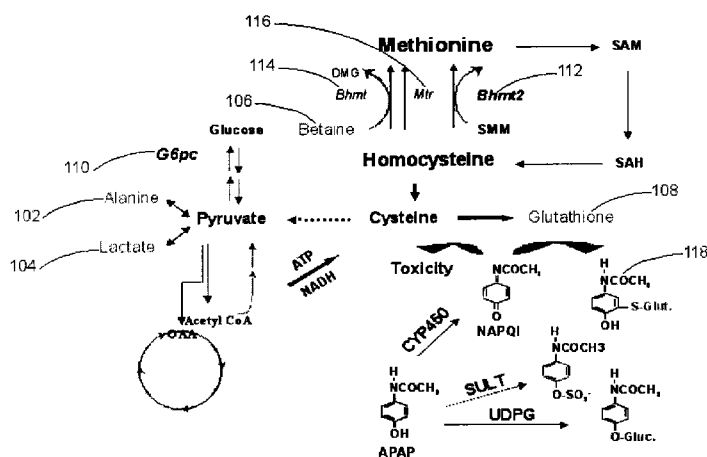


Figure 1

(57) Abstract: The present disclosure relates generally to a co-formulation or co-administration of acetaminophen with one or more chemicals that are metabolites or their derivatives in the methionine and glutathione biosynthesis pathways, for protection against acetaminophen-induced liver toxicity. The formulation may comprise acetaminophen and diet supplements such as S-methylmethionine (SMM). The present disclosure also encompasses methods of providing pain or fever relief with protection against acetaminophen-induced liver injury with the formulations described herein. Embodiments of the present disclosure also include co-formulation or co-administration of an agent and one or more other chemicals for reducing agent-induced liver toxicity via depletion of glutathione.

— *before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments (Rule 48.2(h))*

(88) Date of publication of the international search report:
28 July 2011

INTERNATIONAL SEARCH REPORT

International application No.
PCT/US2010/049093**A. CLASSIFICATION OF SUBJECT MATTER****A61K 31/185(2006.01)i, A61K 31/195(2006.01)i, A61K 31/4015(2006.01)i, A61K 31/4025(2006.01)i, A61K 31/506(2006.01)i, A61P 1/16(2006.01)i**

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)

A61K 31/185; A61K 31/195; C07J 9/00; A61K 31/205; A61K 31/727; A61P 25/08; A61K 31/40

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Korean utility models and applications for utility models

Japanese utility models and applications for utility models

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)

eKOMPASS(KIPO internal) & Keywords: acetaminophen, S-adenosylhomocysteine(SAH), S-methylmethionine(SMM), cystine, betaine

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	US 2008-0200533 A1 (KRISHNAN, RAMU) 21 August 2008 See claims 1, 2, 12, 16, 17	1-19, 27-32, 35
A	US 2006-0128657 A1 (MESSADEK, JALLAL) 15 June 2006 See claims 1, 45	1-19, 27-32, 35
A	US 05428063 A (BARAK, ANTHONY J. et al.) 27 June 1995 See claim 8	1-19, 27-32, 35
A	EP 2011491 A1 (INSTITUT NATIONAL DE LA SANTE ET DE LA RECHERCHE MEDICALE (INSERM)) 07 January 2009 See claims 1, 2	1-19, 27-32, 35

 Further documents are listed in the continuation of Box C. See patent family 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 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 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

27 MAY 2011 (27.05.2011)

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Name and mailing address of the ISA/KR

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INTERNATIONAL SEARCH REPORT

International application No.

PCT/US2010/049093**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.: 20-26, 33, 34
because they relate to subject matter not required to be searched by this Authority, namely:
Claims 20-26, 33, 34 pertain to methods for treatment of the human body by therapy/diagnostic methods, and thus relate to a subject matter which this International Searching Authority is not required, under Article 17(2)(a)(i) of the PCT and Rule 39.1(iv) of the Regulation under the PCT, to search.
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:

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 fee, this Authority did not invite payment of any additional fee.
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.:
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.

INTERNATIONAL SEARCH REPORT

Information on patent family members

International application No.

PCT/US2010/049093

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