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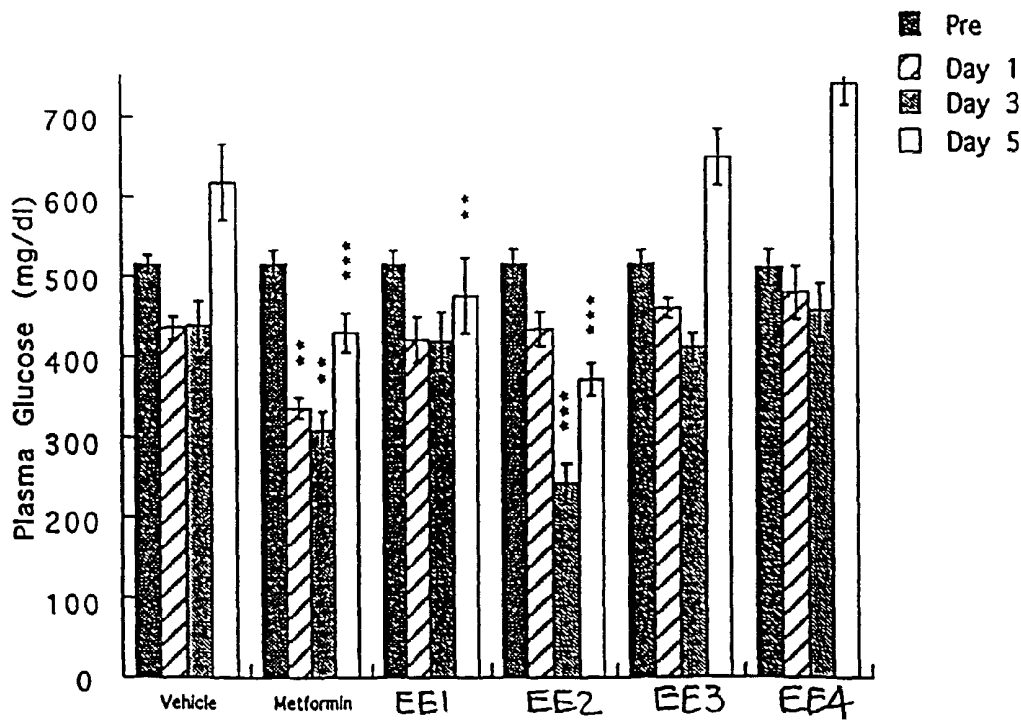
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(54) Title: COMPOSITIONS CONTAINING HYPOGLYCEMICALLY ACTIVE STILBENOID



(57) Abstract: The use of isolated or purified stilbenoid compounds including longistyline A-2-carboxylic acid as hypoglycemic agents or to lower serum glucose levels is described. The invention also relates to the use of such stibenoid compounds in combination with other hypoglycemic agents.

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For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

INTERNATIONAL SEARCH REPORT

International Application No
PCT/US 01/25382

A. CLASSIFICATION OF SUBJECT MATTER
 IPC 7 A61K31/192 A61K31/05 A61K31/085 A61P3/10

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 A61K

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, WPI Data, PAJ, CHEM ABS Data, EMBASE, SCISEARCH, MEDLINE, NAPRALERT BIOSIS

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WO 00 26167 A (BERNARDON JEAN MICHEL ;GALDERMA RES & DEV (FR)) 11 May 2000 (2000-05-11) abstract page 1, line 4 - line 7 page 2, line 11 -page 5, line 2 page 15, line 15 - line 25 page 16, line 24 -page 17, line 8 claims 1,2,9-12 --- -/--	1,6,7, 11,12

Further documents are listed in the continuation of box C.

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

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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	DATABASE CA 'Online! CHEMICAL ABSTRACTS SERVICE, COLUMBUS, OHIO, US; COTIAS, CLAUDIO TENORIO ET AL: "Longistyliline C, antibiotic isolated from Lonchocarpus longistyllus. Preliminary results of its pharmacological properties" retrieved from STN Database accession no. 93:37284 XP002195757 abstract & REV. QUIM. IND. (RIO DE JANEIRO) (1979), 48(564), 12-15,	1,2,4,5
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X	----- ASAKAWA Y ET AL: "PRENYL BIBENZYLs FROM THE LIVERWORTS RADULA-PERROTTETII AND RADULA-COMPLANATA" PHYTOCHEMISTRY (OXFORD), vol. 30, no. 1, 1991, pages 235-252, XP008001857 ISSN: 0031-9422 abstract examples CMPDS10-21,24,25 page 244, column 1, paragraph 3 -column 2, paragraph 1 tables 3,4 page 250, column 2, paragraph 4	1-5
X	----- ASAKAWA, YOSHINORI ET AL: "Chemosystematics of bryophytes. Part 39. Prenyl bibenzyls from the liverwort Radula kojana" PHYTOCHEMISTRY (1991), 30(1), 219-34, XP008001856 abstract page 219, column 1, paragraph 1 examples CMPDS1-8 page 227, column 2, paragraph 3 page 233, column 2, paragraph 5 ----- -/--	1,2,4,5

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Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
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X	MITSCHER L A ET AL: "ANTI MICROBIAL AGENTS FROM HIGHER PLANTS PRENYLATED FLAVONOIDS AND OTHER PHENOLS FROM GLYCYRRHIZA-LEPIDOTA" PHYTOCHEMISTRY (OXFORD), vol. 22, no. 2, 1983, pages 573-576, XP008001858 ISSN: 0031-9422 abstract page 573, column 1, paragraph 1 - paragraph 3 example CMPD1 table 1 ---	1-3,5
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International Application No
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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.
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A	ESPOSITO AVELLA M ET AL: "Evaluation of traditional medicine: effects of Cajanus cajan L. and of Cassia fistula L. on carbohydrate metabolism in mice!. Evaluacion de la medicina tradicional: efectos de Cajanus cajan L. (Guandu) y de Cassia fistula L. (canafistula) en el metabolismo de los carbohidratos en el raton." REVISTA MEDICA DE PANAMA, (1991 JAN) 16 (1) 39-45., XP008001759 page 44, column 1, paragraph 3 -column 2, paragraph 1	6-16
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A	WO 99 56737 A (INST NAT SANTE RECH MED ;MILGROM EDWIN (FR); CASPER ROBERT FREDERI) 11 November 1999 (1999-11-11) page 1, line 5 - line 9 page 5, line 7 - line 12 page 5, line 19 -page 6, line 4 page 7, line 20 - line 23 page 8, line 16 - line 19 page 14, line 23 -page 15, line 4 page 15, line 25 -page 17, line 2 page 18, line 19 - line 24 claims 1-4,8,10,14,19	1-16

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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	DATABASE NAPRALERT 'Online! CHEN D.H., LI H.Y., LIN H.: "Chemical constituents of Cajanus Cajan (L) millisp. leaves" Database accession no. 95:6363 XP002195759 abstract & CHEN D.H., LI H.Y., LIN H.: "Chemical constituents of Cajanus Cajan (L.) millisp. leaves" CHUNG TS'AO YAO, vol. 18, no. 10, 1985, pages 2-7, ---	10
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P,A	WO 00 69430 A (NAG BISHWAJIT ;CALYX THERAPEUTICS INC (US); DEY DEBENDRANATH (US);) 23 November 2000 (2000-11-23) cited in the application abstract page 3, line 16 -page 4, line 12 page 9, line 1 -page 10, line 9 claims 13,14,19 ---	1-16
L	WO 02 13811 A (INSMED CORP) 21 February 2002 (2002-02-21) the whole document Priority ---	1-16
L	WO 02 13809 A (INMAN WAYNE D ;HOPP DAVID CRAIG (US)) 21 February 2002 (2002-02-21) Priority page 5, paragraph 5 -page 6, paragraph 7 page 9, paragraph 1 -page 11, paragraph 3 examples CMPDSA,I page 14, paragraph 2 - paragraph 3 page 22, paragraph 2 -page 23, paragraph 2 page 24, paragraph 4 -page 25, paragraph 4 page 27, paragraph 5 -page 28, paragraph 2 claims 1-9,11,12,15,21,22 ---	1-16
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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.
L	WO 02 13806 A (HOPPE DAVID CRAIG ; INMAN WAYNE D (US)) 21 February 2002 (2002-02-21) Priority page 14, paragraph 2 - paragraph 3 page 22, paragraph 2 - page 23, paragraph 2 page 24, paragraph 4 - page 25, paragraph 4 page 27, paragraph 5 - page 28, paragraph 2 claims 1,9,11,12,17-20 page 5, paragraph 5 - page 6, paragraph 7 page 9, paragraph 1 - page 11, paragraph 3 examples CMPDSA, I -----	1-16

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.2

Present claim 14 relates to a rather large number of possible compounds, namely "a biguanide", "a sulfonylurea", "a PPARgamma agonist", "an alpha-glucosidase inhibitor", "a beta-adrenoceptor agonist". Support within the meaning of Article 6 PCT and/or disclosure within the meaning of Article 5 PCT is to be found, however, for only a very small proportion of the compounds claimed. In the present case, the claims so lack support, and the application so lacks disclosure, that a meaningful search over the whole of the claimed scope is impossible. Consequently, the search has been carried out for those parts of the claims which appear to be supported and disclosed, namely those parts relating to the compounds specifically mentioned in the description p. 31, lines 7-15.

The applicant's attention is drawn to the fact that claims, or parts of claims, relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure.

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Box I Observations where certain claims were found unsearchable (Continuation of item 1 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:

Although claims 6-16 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
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:

see FURTHER INFORMATION sheet PCT/ISA/210
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 II Observations where unity of invention is lacking (Continuation of item 2 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.
- No protest accompanied the payment of additional search fees.

INTERNATIONAL SEARCH REPORT

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