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LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW,
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*For two-letter codes and other abbreviations, refer to the "Guid-
ance Notes on Codes and Abbreviations" appearing at the begin-
ning of each regular issue of the PCT Gazette.*

(54) Title: UROTENSIN-II AGONISTS AND ANTAGONISTS

(57) Abstract: The present invention features a novel class of cyclic polypeptides that have U-II antagonist and agonist activity. The invention also features methods for treating physiological or psychological conditions characterized by an excess or under expression of Ur4otensin-II.



WO 02/032932 A3

INTERNATIONAL SEARCH REPORT

International Application No

PCT/US 01/50724

A. CLASSIFICATION OF SUBJECT MATTER

IPC 7 C07K14/575 A61K38/22

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 C07K 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, BIOSIS, 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	HOCART SJ ET AL.: "Highly Potent Cyclic Disulfide Antagonists of Somatostatin" JOURNAL OF MEDICINAL CHEMISTRY, vol. 42, no. 11, 3 June 1999 (1999-06-03), pages 1863-1871, XP002189091 the entire document, in particular tables 1 and 2, page 1864, left-hand column and page 1868, right-hand column ---	1-3,5, 8-10,12, 15-23
X	HOCART SJ ET AL.: "Potent Antagonists of Somatostatin: Synthesis and Biology" JOURNAL OF MEDICINAL CHEMISTRY, vol. 41, 26 March 1998 (1998-03-26), pages 1146-1154, XP000749590 - page 1146; tables 1,2 --- -/--	1-3,5, 8-10,12, 15-23



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

Date of the actual completion of the international search

20 November 2002

Date of mailing of the international search report

29/11/2002

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INTERNATIONAL SEARCH REPORT

International Application No
PCT/US 01/50724

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	NISHIKI M ET AL.: "Histopathological Improvement of Acromegalic Cardiomyopathy by Intermittent Subcutaneous Infusion of Octreotide" ENDOCRINE JOURNAL, vol. 44, no. 5, 1997, pages 655-660, XP008010820 abstract ----	16-23
A	NEWBY DE & JALAN R: "Urotensin II: Better than somatostatin for portal hypertension?" HEPATOLOGY, vol. 31, no. 5, May 2000 (2000-05), pages 1201-1202, XP008010830 page 1201 ----	16-23
A	FR 2 786 489 A (INSTITUT NATIONAL DE LA SANTE ET DE LA RECHERCHE MEDICALE INSERM) 2 June 2000 (2000-06-02) the entire document, in particular SEQ ID NOs 3 and 9, page 8 and figure 1 -----	1-27

INTERNATIONAL SEARCH REPORT

International application No.
PCT/US 01/50724

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 16-23 and 25-27 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.: 1-27 (all partially)
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.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.2

Claims Nos.: 1-27 (all partially)

The scope of claims 1-27, in as far as the expression "variant" is concerned, is so unclear (Article 6 PCT) that a meaningful search was impossible with regard to this expression.

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 polypeptides as such within the meaning of claim 1.

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.

INTERNATIONAL SEARCH REPORT

information on patent family members

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Patent document cited in search report		Publication date	Patent family member(s)	Publication date
FR 2786489	A	02-06-2000	FR 2786489 A1	02-06-2000
			EP 1131436 A1	12-09-2001
			WO 0031265 A1	02-06-2000
			JP 2002530110 T	17-09-2002