SUBSTITUTED GAMMA LACTAMS AS THERAPEUTIC AGENTS

Abstract: Use of a compound of Formula (I) or a pharmaceutically acceptable salt, prodrug, or a metabolite thereof is disclosed herein, where Y, A, and B are as described herein, for use as medicaments in the treatment of the inflammatory bowel disease or glaucoma. Some compounds of Formula (I) are new.
**INTERNATIONAL SEARCH REPORT**

**International application No**
PCT/US2006/007797

**A. CLASSIFICATION OF SUBJECT MATTER**

INV. C07D207/26 C07D405/04 C07D409/04 A61K31/4015 A61K31/402
A61K31/4025 A61P27/06 A61P29/00

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 C07D

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, CHEM ABS Data, BIOSIS

**C. DOCUMENTS CONSIDERED TO BE RELEVANT**

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<tr>
<th>Category</th>
<th>Citation of document, with indication, where appropriate, of the relevant passages</th>
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<td>A</td>
<td>WO 03/103604 A (APPLIED RESEARCH SYSTEMS ARS HOLDING N.V.; ARALDI; GIAN, LUCA; REDDY, A) 18 December 2003 (2003-12-18) claims 1,25 examples 1-80 table 1</td>
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<td>A</td>
<td>WO 2004/037813 A (MERCK FROSST CANADA &amp; CO; BILLOT, ZAVIER; BEUNARD; JEAN-LUC; HAN, YONG) 6 May 2004 (2004-05-06) the whole document</td>
<td>1-15</td>
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**Date of the actual completion of the international search**

6 July 2006

**Date of mailing of the international search report**

01/09/2006

**Name and mailing address of the ISA/ European Patent Office, P.B. 3818 Patentlaain 2 NL - 2280 HV Rijswijk, Tel. (+31-70) 340-2040, Tx. 31 651 eipo nl, Fax: (+31-70) 340-3016**

**Authorized officer**

Giacobbe, S
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<td>A</td>
<td>US 6 573 294 B1 (OLD DAVID W ET AL) 3 June 2003 (2003-06-03) claims 1,7</td>
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INTERNATIONAL SEARCH REPORT

Box 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. [x] 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 8.4(a).

Box 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.

[ ] No protest accompanied the payment of additional search fees.
Continuation of Box II.2

Claims Nos.: –

The present independent claim 1 relates to an extremely large number of possible compounds. Support and disclosure in the sense of Article 6 and 5 PCT is to be found however for only a very small proportion of them, see Table starting on p. 68. In particular all the compounds present in this Table fall under the much narrower scope of the Markush formula of present claim 10, at least as far as those having a measurable activity are concerned. The non-compliance with the substantive provisions is such that the search was performed taking into consideration the non-compliance in determining its extent (PCT Guidelines 9.19 and 9.23).

The search was therefore restricted to the general Formula of claim 10 and to the specific compounds of claim 13 as such and to their use in the treatment of glaucoma and inflammatory bowel disease.

The applicant’s attention is drawn to the fact that 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. If the application proceeds into the regional phase before the EPO, the applicant is reminded that a search may be carried out during examination before the EPO (see EPO Guideline C-VI, 8.5), should the problems which led to the Article 17(2) declaration be overcome.
<table>
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