Title: RUFOMYCINS AND DERIVATIVES THEREOF USEFUL AS INHIBITORS OF MULTI-DRUG RESISTANCE ASSOCIATED PROTEIN-1 (MRP-1)

Abstract: The present invention provides rufomycin factors and derivatives thereof useful as inhibitors of multi-drug resistant protein. As such these compounds are useful in combination therapy for the treatment of multi-drug resistant diseases. The present invention also provides novel rufomycin factors and derivatives thereof and their pharmaceutical compositions.
INTERNATIONAL SEARCH REPORT

A. CLASSIFICATION OF SUBJECT MATTER
IPC 7 C07K/64 A61K38/00

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)
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, CHEM ABS Data, BIOSIS, EMBASE, WPI Data

C. DOCUMENTS CONSIDERED TO BE RELEVANT

<table>
<thead>
<tr>
<th>Category</th>
<th>Citation of document, with indication, where appropriate, of the relevant passages</th>
<th>Relevant to claim No.</th>
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<tbody>
<tr>
<td>A</td>
<td>US 3 655 879 A (NAKAZAWA KOITI ET AL) 11 April 1972 (1972-04-11)</td>
<td>14</td>
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<td>A</td>
<td>FUJINO M. ET AL.: &quot;Tryptophan Moiety of Rufomycin Homologs&quot; CHEM.PHARM.BULL., vol. 12, no. 11, 1964, pages 1390-2, XP000945345 figure 1</td>
<td></td>
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</table>

Further documents are listed in the continuation of box C.

Patent family members are listed in annex.

Date of the actual completion of the international search
3 January 2001

Date of mailing of the international search report
8.01.01

Name and mailing address of the ISA
European Patents Office, P.B. 5018 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016

Authorized officer
Deffner, C-A
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<th>Citation of document, with indication, where appropriate, of the relevant passages</th>
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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. **X** Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
   
   Although claims 1 to 13 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:

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.
<table>
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<tr>
<td>US 3655879 A</td>
<td>11-04-1972</td>
<td>CH 401357 A</td>
<td>31-10-1965</td>
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