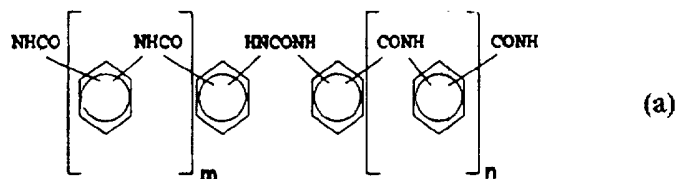
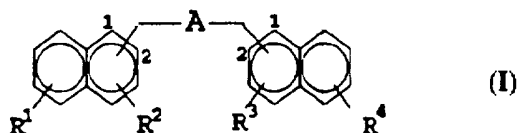


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(54) Title: USE OF DINAPHTHALENES COMPOUNDS AS ANTIPROLIFERATIVE AGENTS



(57) Abstract

The use of a compound with structural formula (I), wherein R^1 to R^4 are each independently one or more of -X, -N₃, -NO₂, halo, trifluoromethyl, R⁵, OR⁵, -CH₂OR⁵, -OCOR⁵, -CH₂OCOR⁵, -NHCOR⁵, -CH₂NHCOR⁵, -NR⁵R⁶, -CH₂NR⁵R⁶, -CH₂NO₂, CONR⁵R⁶, CH₂CONR⁵R⁶, -COOR⁵, -CH₂COOR⁵, -CHO and -CH₂CHO and -X is independently -SO₃R⁵, -CH₂PO₃R⁵R⁶, -CH₂SO₃R⁵, -OSO₃R⁵, -CH₂OSO₃R⁵, -NHSO₃R⁵, -CH₂NHSO₃R⁵, -OPO₃R⁵R⁶, -CH₂OPO₃R⁵R⁶ and -PO₃R⁵R⁶ where R⁵ and R⁶ are chosen independently from -H and lower alkyl and wherein A is a chemical group comprising between 5 and 30 bonds directly linking the naphthyl groups provided that (i) the compound is not suramin and (ii) when A is not (a), wherein m and n are independently 0, 1 or 2, then at least one of R^1 to R^4 is -OH or an acidic group or a pharmaceutically acceptable salt, ester, salt of such ester or amide of such compounds, in the manufacture of a medicament for use in reducing undesired angiogenesis, treating cancer, treating fibrotic disease, or treating diseases benefiting from antagonism of the action of fibroblast, vascular endothelial and transforming growth factors. Preferably A is (a) and m and n are independently 0, 1 or 2. Novel pharmaceutical compositions with other medicaments are provided.

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A. CLASSIFICATION OF SUBJECT MATTER

IPC 5 A61K31/165 A61K31/17 A61K31/18 A61K31/00 A61K31/185

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 5 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)

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WO,A,91 10649 (FARMITALIA) 25 July 1991 * see claims 1-4,8 and page 11, lines 1-20 ---	1,7-9, 45,60,61
X	WO,A,90 15816 (UPJOHN) 27 December 1990 cited in the application * see page 4 lines 4-27, page 7 lines 27-35, claims 1-3,6,13,14 ---	1,45,60
A	WO,A,92 05153 (AGOURON) 2 April 1992 *see claims 1,42,45 * ---	1,7-26, 45,46, 60,61
A	EP,A,0 354 818 (HADASSAH) 14 February 1990 * see Claims 1 and 15; Example 4; page 3 line 35 - page 6 line 12 * -----	1



Further documents are listed in the continuation of box C.



Patent family members are listed in annex.

* Special categories of cited documents :

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Date of the actual completion of the international search

28 March 1994

Date of mailing of the international search report

17. 06. 94

Name and mailing address of the ISA

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INSERT B.

INTERNATIONAL SEARCH REPORT

Inter. onal application No.

PCT/GB93/02493

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:
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:

For further information please see form PCT/ISA/206 dated 21.04.1994.

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.:

1,45,46,60,61 and 7-26 partially

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

Information on patent family members

International Application No

PC 1/GB 93/02493

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
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