Title: METHOD OF MODULATING VASCULARIZATION

Abstract: A method of modulating vascularization in a tissue of a mammal comprises controlling a PAR signaling pathway (e.g., the PAR-1 or PAR-2 signaling pathway) in a mammalian tissue, for example, by controlling phosphorylation of tissue factor cytoplasmic domain (i.e., phosphorylation of Ser258 of the cytoplasmic tail of TF). In a preferred method pathological neovascularization in a mammal is treated by administering to a mammal suffering from pathological neovascularization, a therapeutically effective amount of a PAR signaling pathway inhibitor. Preferably the mammal is a human.
## INTERNATIONAL SEARCH REPORT

### A. CLASSIFICATION OF SUBJECT MATTER
- **IPC:** A61K 38/00 (2006.01), 39/00 (2006.01), 39/395 (2006.01), 48/00 (2006.01); C07H 21/04 (2006.01)
- **USPC:** 424/145.1, 146.1, 184.1; 514/2, 44; 536/24.5

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)
- **U.S.:** 424/145.1, 146.1, 184.1; 514/2, 44; 536/24.5

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 practicable, search terms used)
Please See Continuation Sheet

### 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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</table>

**X** Further documents are listed in the continuation of Box C.  
**See patent family annex.**

- **"A"** 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 application or patent 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

Date of the actual completion of the international search: 19 March 2007 (19.03.2007)

Date of mailing of the international search report: 24 April 2007

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<tr>
<td>X</td>
<td>US 5,976,534 A (HART et al.) 02 November 1999 (02.11.1999) (see entire document).</td>
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<td>X, E</td>
<td>US 7,009,036 B2 (FACCHIANO et al.) 07 March 2006 (07.03.2006) (see entire document).</td>
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<tr>
<td>X</td>
<td>SEBITI et al. Design of growth factor antagonists with antiangiogenic and antitumor properties. Oncogene. 2000, Vol. 19, pages 6566-6573 (see entire document).</td>
<td>1, 5, 8, 9</td>
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Continuation of B. FIELDS SEARCHED Item 3:
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