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(54) Title: METHODS FOR ALZHEIMER’S DISEASE TREATMENT AND COGNITIVE ENHANCEMENT

(57) Abstract: The present invention relates to compositions and methods to modulate e-secretase and/or to improve cognitive ability. The invention further relates the improved/enhanced cognitive ability in diseased individuals, particularly Alzheimer’s Disease patients, and treatment thereof through increased sAPP production. Macrocyclic lactones (i.e. bryostatin class and neristatin class) are compounds preferred for use with the present composition. The present invention also provides methods for increasing the generation of non-amyloidogenic soluble APP comprising the activation of protein kinase C (PKC) by administering an effective amount of PKC activator(s).
INTERNATIONAL SEARCH REPORT

A. CLASSIFICATION OF SUBJECT MATTER
   IPC(7) : A61k 31/35, 31/335
   US CL : 514/450, 451, 449
   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. : 514/450, 451, 449
   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)
   CAS ONLINE MEDLINE BIOSIS search terms: bryostatin, neristatin, PKC activator, Alzheimer, dementia

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>Y</td>
<td>US 6,880,784 A (DRIEDGER et al) 27 June 2000 (27.06.2000), column 2, line 47 to column 3, line 4, and the claims.</td>
<td>1-36</td>
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Further documents are listed in the continuation of Box C. See patent family annex.

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