



## INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

|                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 |           |                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    |
|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <b>(51) International Patent Classification <sup>7</sup>:</b><br><b>C07D 333/00</b>                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             | <b>A2</b> | <b>(11) International Publication Number:</b> <b>WO 00/44740</b><br><b>(43) International Publication Date:</b> 3 August 2000 (03.08.00)                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           |
| <b>(21) International Application Number:</b> PCT/US00/01980<br><b>(22) International Filing Date:</b> 27 January 2000 (27.01.00)<br><b>(30) Priority Data:</b><br>09/239,091 27 January 1999 (27.01.99) US<br><b>(71) Applicant:</b> AMERICAN CYANAMID COMPANY [US/US];<br>Five Giralda Farms, Madison, NJ 07940-0874 (US).<br><b>(72) Inventors:</b> LEVIN, Jeremy, Ian; 19 Long Meadow Drive,<br>New City, NY 10956 (US). CHEN, James, Ming; 7<br>Sergeant David Stoddard Court, Bedminster, NJ 07921 (US).<br>NELSON, Frances, Christy; 69 Terrace Heights, Wyckoff,<br>NJ 07481 (US).<br><b>(74) Agents:</b> BARRETT, Rebecca, R.; American Home Products<br>Corporation, Patent Law Department -2B, One Campus<br>Drive, Parsippany, NJ 07054 (US) et al. |           | <b>(81) Designated States:</b> AE, AL, AM, AT, AU, AZ, BA, BB, BG,<br>BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE,<br>ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP,<br>KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA,<br>MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU,<br>SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG,<br>UZ, VN, YU, ZA, ZW, ARIPO patent (GH, GM, KE, LS,<br>MW, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ,<br>BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE,<br>CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC,<br>NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA,<br>GN, GW, ML, MR, NE, SN, TD, TG).<br><br><b>Published</b><br><i>Without international search report and to be republished<br/>         upon receipt of that report.</i> |
| <b>(54) Title:</b> HETEROARYL ACETYLENIC SULFONAMIDE AND PHOSPHINIC ACID AMIDE HYDROXAMIC ACID TACE<br>INHIBITORS<br><br><div style="text-align: center;"> <p style="text-align: right;">(I)</p> </div><br><b>(57) Abstract</b><br><p>Compounds of formula (I) are useful in treating disease conditions mediated by TNF-<math>\alpha</math> such as rheumatoid arthritis, osteoarthritis, sepsis, AIDS, ulcerative colitis, multiple sclerosis, Crohn's disease and degenerative cartilage loss.</p>                                                                                                                                                                                                                                                           |           |                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    |