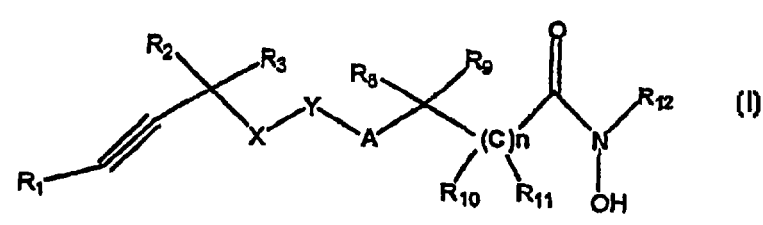




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(21) International Application Number: PCT/US00/01864 (22) International Filing Date: 27 January 2000 (27.01.00) (30) Priority Data: 09/238,038 27 January 1999 (27.01.99) US (71) Applicant: AMERICAN CYANAMID COMPANY [US/US]; Five Giralda Farms, Madison, NJ 07940-0874 (US). (72) Inventors: LEVIN, Jeremy, Ian; 19 Long Meadow Drive, New City, NY 10956 (US). VENKATESAN, Aranapakam, Mudumbai; 97-07, 63rd Road #9K, Rego Park, NY 11374 (US). CHEN, James, Ming; 7 Sergeant David Stoddard Court, Bedminster, NJ 07921 (US). ZASK, Arie; 21 East 90 Street, New York, NY 10128 (US). SANDANAYAKA, Vincent, Premarana; #1 Earl Court, Wesley Hills, NY 10952 (US). DU, Mila, Ti; 17 Heather Hill Lane, Suffern, NY 10901 (US). BAKER, Jannie, Lea; 127 Rockinchair Road, White Plains, NY 10607 (US). (74) Agents: BARRETT, Rebecca, R.; American Home Products Corporation, Patent Law Dept. - 2B, One Campus Drive, Parsippany, NJ 07054 (US) et al.		(81) Designated States: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG). Published <i>With international search report.</i> <i>Before the expiration of the time limit for amending the</i> <i>claims and to be republished in the event of the receipt of</i> <i>amendments.</i>  *G00001*
(54) Title: ALKYNYL CONTAINING HYDROXAMIC ACID DERIVATIVES, THEIR PREPARATION AND THEIR USE AS MATRIX METALLOPROTEINASE (MMP) INHIBITORS/TNF-ALPHA CONVERTING ENZYME (TACE) INHIBITORS		
(57) Abstract Compounds of formula (I) wherein R ₁ is hydrogen, aryl, heteroaryl, alkyl of 1-8 carbon atoms, alkenyl of 2-6 carbon atoms, alkynyl of 2-6 carbon atoms, cycloalkyl of 3-6 carbon atoms, or -C ₄ -C ₈ -cycloheteroalkyl; R ₂ and R ₃ are hydrogen, alkyl of 1-6 carbon atoms, -CN, or CCH; R ₈ , R ₉ , R ₁₀ and R ₁₁ are hydrogen, aryl or heteroaryl, cycloalkyl of 3-6 carbon atoms, -C ₄ -C ₈ -cycloheteroalkyl, alkyl of 1-18 carbon atoms, alkenyl of 2-18 carbon atoms, alkynyl of 2-18 carbon atoms; with the proviso that one of the pairs R ₈ and R ₉ , R ₉ and R ₁₀ or R ₁₀ and R ₁₁ , together with the carbon atom or atoms to which they are attached, form a cycloalkyl ring of 3-6 carbon atoms, or a -C ₄ -C ₈ -cycloheteroalkyl ring; R ₁₂ is hydrogen, aryl or heteroaryl, cycloalkyl of 3-6 carbon atoms; -C ₄ -C ₈ -cycloheteroalkyl, or alkyl of 1-6 carbon atoms; A, X are O, S, SO, SO ₂ , NR ₇ , or CH ₂ ; Y is aryl or heteroaryl, with the proviso that A and X are not bonded to adjacent atoms of Y, and n is 0-2; or a pharmaceutically acceptable salt thereof; useful in the treatment of arthritis, tumor metastasis, tissue ulceration, abnormal wound healing, periodontal disease, bone disease, diabetes (insulin resistance) and HIV infection.  (I)  *162162*		