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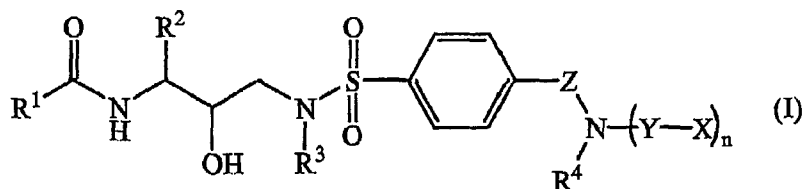
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(54) Title: HIV PRODRUGS CLEAVABLE BY CD26



(57) Abstract: The present invention provides new prodrugs which are conjugates of a therapeutic compound and a peptide wherein the conjugate is cleavable by dipeptidyl-peptidases, more preferably by CD26, also known as DPPIV (dipeptidyl aminodipeptidase IV).

The present prodrugs have the formula (I), the stereoisomeric forms and salts thereof, wherein n is 1 to 5; Y is proline, alanine, hydroxyproline, dihydroxyproline, thiazolidinecarboxylic acid (thioprolin), dehydroproline, pipercolic acid (L-homoprolin), azetidinecarboxylic acid, aziridinecarboxylic acid, glycine, serine, valine, leucine, isoleucine and threonine; X is selected from any amino acid in the D- or L-configuration; X and Y in each repeat of [Y-X] are chosen independently from one another and independently from other repeats; Z is a direct bond or a bivalent straight or branched saturated hydrocarbon group having from 1 to 4 carbon atoms; R¹ is an aryl, heteroaryl, aryloxy, heteroaryloxy, aryloxyC₁₋₄alkyl, heterocycloalkyloxy, heterocycloalkylC₁₋₄alkoxy, heteroaryloxyC₁₋₄alkyl, heteroarylC₁₋₄alkyloxy; R² is arylC₁₋₄alkyl; R³ is C₁₋₁₀alkyl, C₂₋₆alkenyl or C₃₋₇cycloalkylC₁₋₄alkyl; R⁴ is hydrogen or C₁₋₄alkyl. The present invention furthermore provides the use of said prodrugs as medicines as well as a method of producing said prodrugs.



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