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(54) Title: COMPOSITIONS AND METHODS FOR MODULATING PHYSIOLOGY OF EPITHELIAL JUNCTIONAL ADHESION MOLECULES FOR ENHANCED MUCOSAL DELIVERY OF THERAPEUTIC COMPOUNDS

Human JAM-1: 299 amino acids.

MGTKAQVERKLLCLFILAILLCSLALGSVTVHSSEPEVRIPENNPVKLSCAYSGFSSP RVEWKFDQGDTTRLVCYNNKITASYEDRVTFLPTGITFKSVTREDTGTYTCMVS EEGGNSYGEVKVKLIVLVPPSKPTVNIPSSATIGNRAVLTCSEQDGSPPSEYTWF KDGIVMPTNPKSTRAFSNSSYVLNPTTGELVFDPLSASDTGEYSCEARNGYGTPM TSNAVRMEAVERNVGVIVAAVLVTLILLGILVFGIWFAYSRGHFDRTKKGTSSKKVI YSQPSARSEGEFKQTSSFLV (SEQ ID NO: 887)

(57) Abstract: Compositions and methods are provided that include a biologically active agent and a permeabilizing agent effective to enhance mucosal delivery of the biologically active agent in a mammalian subject. The permeabilizing agent reversibly enhances mucosal epithelial paracellular transport, typically by modulating epithelial junctional structure and/or physiology at a mucosal epithelial surface in the subject. This effect typically involves inhibition by the permeabilizing agent of homotypic or heterotypic binding between epithelial membrane adhesive proteins of neighboring epithelial cells. Target proteins for this blockade of homotypic or heterotypic binding can be selected from various related junctional adhesion molecules (JAMs), occludins, or claudins. The permeabilizing agent is typically a peptide or peptide analog or mimetic, often selected or derived from an extracellular domain of a mammalian JAM, occludin or claudin protein.





INTERNATIONAL SEARCH REPORT

International application No.

PCT/US03/19994

A. CLASSIFICATION OF SUBJECT MATTER IPC(7) : A61K 38/00				
US CL : 514/12				
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.: 514/12, 2; 530/324				
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				
Category *	Citation of document, with indication, where appro		Relevant to claim No.	
A	AURRAND-LIONS et al. JAM-2, a Novel Immunoglob Expressed by Endothelial and Lymphatic Cells. Journal 2001, Vol. 276, No. 4, pages 2733-2741. See entire do	Biological Chemistry. 26 January	1-92	
Y	US 6,110,747 A (BLASCHUK et al.) 29 August 2000 (29.08.2001), see abstract.	1	
Α			2-92	
X US 6,248,864 B1 (BLASCHUK et al.) 19 June 200		9.06.2001), see abstract, and	1	
Α	columns 57-58.		2-92	
Further	documents are listed in the continuation of Box C.	See patent family annex.		
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"A" document	defining the general state of the art which is not considered to be lar relevance	date and not in conflict with the applica principle or theory underlying the inver		
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"O" document	referring to an oral disclosure, use, exhibition or other means	combined with one or more other such being obvious to a person skilled in the		
Date of the a	Date of the actual completion of the international search Date of mailing of the international search report			
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Continuation of B. FIELDS SEARCHED Item 3:	
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search terms: JAM-1, JAM-2, JAM-3, immunoglobulin, endothelial, occludin, a	adhesion, cell, permeability, modulte
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