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Published:

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(54) Title: METHODS OF IDENTIFYING ANTI-INFLAMMATORY COMPOUNDS

(57) Abstract: A mammalian C-type lectin receptor type is identified which is shown to bind IgG antibodies or Fc fragments, thus inducing IVIG-related reversal of inflammation associated with various immune disorders. The identification of a DC- SIGN receptor type which interacts with IgG to promote a biological response reducing inflammation associated with immune disorders provides for methods of screening and selecting compounds which may be useful in treating various immune disorders by acting to modulate a DC-SIGN1+1 cell to signal a second effector macrophage, causing an increase in expression of the FcyRIIB receptor and in turn inhibiting a cellular-mediated inflammatory response.

INTERNATIONAL SEARCH REPORT

International application No. PCT/US 09/41441

A. CLASSIFICATION OF SUBJECT MATTER IPC(8) - G01N 33/48 (2009.01) USPC - 435/7.2 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) USPC: 435/7.2				
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched USPC: 435/7.2, 435/4 (text search)				
Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) Electronic data bases: (EPAB, JPAB, PGPB, USPT); Google Scholar Search Terms: IVIG, anti-inflammatory, Fc receptor, sialylated IgG Fc, DC-SIGN (CD209), DC-SIGNR, SIGN-R, SIGN-R1 (CD209a), CLEC4M,				
C. DOCUMENTS CONSIDERED TO BE RELEVANT				
Category*	Citation of document, with indication, where a	ppropriate, of the relevant passages	Relevant to claim No.	
X 	CAPARROS et al. "DC-SIGN ligation on dendritic cells results in ERK and PI3K activation and modulates cytokine production". IN: Blood; 15 May 2006; Vol. 107, No. 10; Pages 3950-8.		1, 2, 4-6, 10, 11	
Y	Especially pg 3951 left col para 2, pg 3954 fig 3e	o, voi. 107, 110. 10, 1 ages 3000-0.	7, 8	
×	REQUENA et al. "Inhibition of HIV-1 transmission in trans from dendritic cells to CD4+ T		13-16, 18, 19	
Y	lymphocytes by natural antibodies to the CRD domain intravenous immunoglobulins." IN: Immunology; April 2 No. 4; Pages 508-518. Especially pg 509 right col para	2008 (Epub 10 November 2007); Vol. 123,	7, 8, 20-22	
Y	ANTHONY et al. "Recapitulation of IVIG anti-inflamma IN: Science; 18 April 2008; Vol. 320, No. 5874; Pages col para 1, pg 374 center col para 2.		20-22	
:				
Further documents are listed in the continuation of Box C.				
* Special categories of cited documents: "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or the				
"E" earlier a	to be of particular relevance "E" earlier application or patent but published on or after the international filing date the principle or theory underlying the invention document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive			
"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other "Y" document of particular relevance; the claimed invention cannot be			claimed invention cannot be	
means	"O" document referring to an oral disclosure, use, exhibition or other combined with one or more other such documents, such combination			
"P" document published prior to the international filing date but later than "&" document member of the same patent family the priority date claimed				
Date of the actual completion of the international search Date of mailing of the international search report				
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Name and mailing address of the ISA/US Mail Stop PCT, Attn: ISA/US, Commissioner for Patents P.O. Box 1450, Alexandria, Virginia 22313-1450 Authorized officer: Lee W. Young				
	0, Аlexandria, Virginia 22313-1450 0. 571-273-3201	PCT Helpdesk: 571-272-4300	ĺ	

INTERNATIONAL SEARCH REPORT

International application No.
PCT/US 09/41441

Box No. II	Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)		
This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:			
	aims Nos.: cause they relate to subject matter not required to be searched by this Authority, namely:		
be	laims Nos.: cause they relate to parts of the international application that do not comply with the prescribed requirements to such an stent that no meaningful international search can be carried out, specifically:		
	laims Nos.: 3, 9, 12, 17, 23, 24 cause they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).		
Box No. III	Observations where unity of invention is lacking (Continuation of item 3 of first sheet)		
This Interna	tional Searching Authority found multiple inventions in this international application, as follows:		
This application contains the following inventions or groups of inventions which are not so linked as to form a single general inventive concept under PCT Rule 13.1. In order for all inventions to be examined, the appropriate additional examination fees must be paid.			
Group I: claims 1, 2, 4-8, 10, 11, 13-16, and 18-22, directed to a method of identifying a test compound useful to activate or suppress anti-inflammatory activity associated with IgG autoantibody-mediated inflammation Group II: claims 25-32, directed to a method of treating an immune disorder comprising administering a compound that binds to and activates a DC-SIGN receptor type or a member of the signal transduction pathway thereof and mediates anti-inflammatory activity, with the proviso that the compound is not IVIG.			
- Please see extra sheet for continuation -			
2. A	s all required additional search fees were timely paid by the applicant, this international search report covers all searchable aims. s all searchable claims could be searched without effort justifying additional fees, this Authority did not invite payment of additional fees. s only some of the required additional search fees were timely paid by the applicant, this international search report covers		
3. L. A	s only some of the required additional search lees were finitely paid by the applicant, this international search report covers ally those claims for which fees were paid, specifically claims Nos.:		
l "" re	o required additional search fees were timely paid by the applicant. Consequently, this international search report is stricted to the invention first mentioned in the claims; it is covered by claims Nos.: 2, 4-8, 10, 11, 13-16, and 18-22		
Remark on	Protest The additional search fees were accompanied by the applicant's protest and, where applicable, the payment of a protest fee. The additional search fees were accompanied by the applicant's protest but the applicable protest fee was not paid within the time limit specified in the invitation. No protest accompanied the payment of additional search fees.		

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US 09/41441 Continuation of Box III: Lack of Unity of Invention The inventions listed as Groups I - II do not relate to a single general inventive concept under PCT Rule 13.1 because, under PCT Rule 13.2, they lack the same or corresponding special technical features for the following reasons: The special technical feature of the Group I claims is a method of identifying a test compound useful to activate or suppress antiinflammatory activity associated with IgG autoantibody-mediated inflammation - not required by the claims of Group II. The special technical feature of the Group II claims is a method of treating an immune disorder comprising administering a compound that binds to and activates a DC-SIGN receptor type or a member of the signal transduction pathway thereof and mediates anti-inflammatory activity, with the proviso that the compound is not IVIG - not required by the claims of Group I. Neither of these special technical features is common to the other group, nor do they correspond to a special technical feature in the other group. The only common technical element shared by the above groups is that they are related to modulation of DC-SIGN to alter anti-inflammatory activity in a subject. This common technical element does not represent an improvement over the prior art of US 2006/0269540 A1 to Robert et al. (see abstract; para [0015] - [0017], [0077], [0083], [0088]). Therefore, the inventions of Group I and Group II lack unity of invention under PCT Rule 13 because they do not share a same or corresponding special technical feature.