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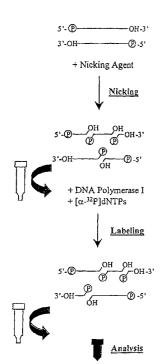
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19 June 2003

[Continued on next page]

(54) Title: DIPEPTIDE SERYL-HISTIDINE AND RELATED OLIGOPEPTIDES CLEAVE DNA, PROTEIN, AND A CARBOXYL ESTER

Nick Translation with Ser-His



Double-stranded DNA substrate (linear \(\DNA \) and circular \(\precept{pBR322} \) tested)

Nicking Reaction (Ser-His or DNase I)

Newly generated 3'OH termini (blue)

Nicked DNA purified on a G-50 Sephadex Quick Spin Column

Labeling Reaction $[\alpha-32P]dNTP + DNA Polymerase I$

Radioactively labeled deoxyribonucleoside monophosphates (red) are incorporated into the DNA

Labeled DNA purified on a G-50 Sephadex Quick Spin Column

(57) Abstract: A compound other than Ser- His, having the structure (Ser/Cys) - Xaa_m - His - Xaa_n (EM'), where 0<=(m+n)<=12, is useful as a nuclease or protease. A compound having the structure EM-L-BM, where EM is EM' above or Ser-His, L is a linker and BM is a nucleotide or amino acid sequence binding moiety, is also useful as a nuclease or protease. The compounds, EM and EM-L-BM which act as nucleases are useful in labeling a nucleic acid by nick translation.

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For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

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A. CLASSII IPC 7	FICATION OF SUBJECT MATTER C07K5/08 C07K5/10 C07K7/06 C07H21/00	C07K7/08	C07K19/00	
According to	International Patent Classification (IPC) or to both national classific	ation and IPC		
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	ata base consulted during the international search (name of data ba BS Data, WPI Data, PAJ, EPO-Interna		erms used)	
C. DOCUME	ENTS CONSIDERED TO BE RELEVANT			
Category °	Citation of document, with indication, where appropriate, of the rel	evant passages	Relevant to claim No.	
X	DATABASE CA 'Online! CHEMICAL ABSTRACTS SERVICE, COLUMOHIO, US; KULIK, WILLEM ET AL: "Fast atom bombardment tandem mass spectrome amino acid sequence determination tripeptides" retrieved from STN Database accession no. 113:6782 (XP002227765 & BIOMEDICAL & ENVIRONMENTAL MASS SPECTROMETRY (1989), 18(10), 910-1989, abstract	etry for n in CA		
X Furt	her documents are listed in the continuation of box C.	Patent family members	s are listed in annex.	
	ategories of cited documents:	*T* later document published aff	ter the international filing date	
consid	ent defining the general state of the art which is not lered to be of particular relevance document but published on or after the international late	cited to understand the prin invention "X" document of particular relev	onflict with the application but notified or theory underlying the name; the claimed invention alor cannot be considered to	
"L" docume which citatio	ent which may throw doubts on priority claim(s) or is cited to establish the publication date of another n or other special reason (as specified)	involve an inventive step w "Y" document of particular relev cannot be considered to in:	then the document is taken alone	
other	ent referring to an oral disclosure, use, exhibition or means ent published prior to the international filing date but han the priority date claimed		eing obvious to a person skilled	
	actual completion of the international search	Date of mailing of the intern	national search report	
1	7 January 2003	04/02/2003		
Name and	mailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2	Authorized officer	Authorized officer	
	Nl. – 2280 HV Rijswijk Tel. (+31–70) 340–2040, Tx. 31 651 epo nl, Fax: (+31–70) 340–3016	Masturzo, P		

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	ation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category °	Citation of document, with indication, where appropriate, of the relevant passages		Relevant to claim No.
X	DATABASE WPI , 1999 Derwent Publications Ltd., London, GB; AN 1999-471871 XP002227766 "New modified amino acid specific enzyme active useful biochemical catalyst" & JP 11 196819 A (M SHIMIZU), 27 July 1999 (1999-07-27) abstract		1
P,X	Y LI ET AL.: "Peptide seryl-histidine and related oligopeptides cleave DNA, protein and a carboxyl ester " BIOORGANIC & MEDICINAL CHEMISTRY., vol. 8, no. 12, December 2000 (2000-12), pages 2675-2680, XP002227764 ELSEVIER SCIENCE LTD., GB ISSN: 0968-0896 the whole document		1-9

nter al application No. PC I/US 01/43079

Box I Observations where certain claims were found unsearchable (Continuation of Item 1 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:
Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
2. X Claims Nos.: 1-9 because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically: see FURTHER INFORMATION sheet PCT/ISA/210
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. No required additional search fees were timely paid by the applicant. Consequently, this international Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.2

Claims Nos.: 1-9

Present claims 1-9 relate to an extremely large number of possible compounds and methods based on these compounds. Support within the meaning of Article 6 PCT and/or disclosure within the meaning of Article 5 PCT is to be found, however, for only a very small proportion of the compounds and methods claimed. In the present case, the claims so lack support, and the application so lacks disclosure, that a meaningful search over the whole of the claimed scope is impossible. Moreover the applicant is informed that the Search Division is not in state to check whether the compounds falling under the wording of claims 1-2 indeed have the claimed nuclease/protease activity. Consequently, the search has been carried out for those parts of the claims which appear to be supported and disclosed, namely those parts relating to the active compounds listed in Table 2 and methods based on them.

The applicant's attention is drawn to the fact that claims, or parts of claims, relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure.

tion on patent family members

Internation pplication No
PCT/US 01/43079

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Patent document cited in search report	Publication date	Patent family member(s)	'	Publication date
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