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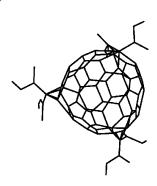
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[Continued on next page]

#### (54) Title: SUBSTITUTED FULLERENES AS MRI CONTRAST AGENTS



(57) Abstract: A contrast agent for enhancing contrast in in vivo magnetic resonance imaging, comprising a water-soluble, non-zero spin isotope-enriched substituted fullerene having a fullerene core or heterofullerene core and at least one substituent group bonded to at least one carbon of the fullerene core. A method of magnetic resonance imaging a sample, comprising (i) administering to the sample a plurality of substituted fullerenes; and (ii) detecting magnetic resonance signals from the sample. In the method, some or all of the plurality of substituted fullerenes may be non-zero spin isotope-enriched, but need not be. In the method, some or all of the plurality of substituted fullerenes may be hyperpolarized, but need not be.

FIG. 1A

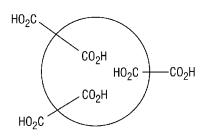


FIG. 1B



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OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

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A. CLASSIFICATION OF SUBJECT MATTER INV. A61K49/10 C01B31/02

According to International Patent Classification (IPC) or to both national classification and IPC

#### **B. FIELDS SEARCHED**

 $\begin{tabular}{ll} \begin{tabular}{ll} Minimum documentation searched & (classification system followed by classification symbols) \\ \begin{tabular}{ll} C01B & \begin{tabular}{ll} \be$ 

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 practical, search terms used)

EPO-Internal, WPI Data, BIOSIS, EMBASE

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Х	US 2003/065206 A1 (BOLSKAR ROBERT D [US] ET AL) 3 April 2003 (2003-04-03)	8,21
′	figure 1, paragraphs 77, 104, 107-116	1–21
X	DE 198 08 865 A1 (BAYERL [DE]; HIRSCH [DE]) 9 September 1999 (1999-09-09)	8,21
Y	Figure 1, column 6 line 65-column 7 line 34.	1-21
•	US 2007/025918 A1 (HURD RALPH E [US]) 1 February 2007 (2007-02-01) Paragraph 22	1-21
,	WO 2007/143558 A1 (UNIV RICE WILLIAM M [US]; RES DEV FOUNDATION [US]; ROSENBLUM MICHAEL G) 13 December 2007 (2007-12-13) Claims 1, 7; Figure 27, page 33 example 5	1-21
	-/	

X Further documents are listed in the continuation of Box C.	See patent family annex.		
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Date of the actual completion of the international search  4 December 2009	Date of mailing of the international search report $14/12/2009$		
Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL – 2280 HV Rijswijk Tel. (+31–70) 340–2040, Fax: (+31–70) 340–3016	Authorized officer  Bettio, Andrea		

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#### INTERNATIONAL SEARCH REPORT

International application No PCT/US2009/037653

C(Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT					
ategory*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.			
Y	BALL G. E. ET AL.: "Structural reassignement of the Mono- and Bis-Addition Products from the Addition Reactions of N-(diphenylmethylene)glycinate Esters to [60]Fullerene under Bingel conditions" J. ORG. CHEM., vol. 70, no. 24, 16 September 2005 (2005-09-16), XP7910747 Page 8572 figure 1, paragraph 1.	1-21			
Y	WO 2005/097807 A2 (LUNA INNOVATIONS INC [US]; GE ZHONGXIN [US]; DORN HARRY C [US]; PHILLI) 20 October 2005 (2005-10-20) Page 8 line 17-page 9 line 7	1-21			
Y	WO 2006/009696 A2 (HEMATROPE PHARMACEUTICALS INC [US]; MELTON LAURA J [US]) 26 January 2006 (2006-01-26) Page 18	1-21			
Y	WO 2007/070466 A2 (HARVARD COLLEGE [US]; MARCUS CHARLES [US]; JOHNSON ALEX [US]; APTEKAR) 21 June 2007 (2007-06-21) Claims 1, 2; page 6 lines 25-30	1-21			
A	MEIER M. ET AL.: "A 13C INADEQUATE and HF-GIAO Study of C60H2 and C60H6 Identification of Ring Currents in a 1,2-Dihydrofullerene"  J. AM. CHEM. SOC., vol. 124, 10 July 2002 (2002-07-10), pages 8090-8094, XP007910752 the whole document	1-21			

1

#### INTERNATIONAL SEARCH REPORT

Information on patent family members

International application No PCT/US2009/037653

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
US 2003065206 A	03-04-2003	US 2003220518 A1	27-11-2003
DE 19808865 A	09-09-1999	NONE	
US 2007025918 A	01-02-2007	WO 2008008075 A2	17-01-2008
WO 2007143558 A	13-12-2007	US 2009214101 A1	27-08-2009
WO 2005097807 A	20-10-2005	US 2008166285 A1	10-07-2008
WO 2006009696 A	26-01-2006	NONE	
WO 2007070466 A	21-06-2007	AU 2006326596 A1 EP 1968442 A2 JP 2009518440 T US 2008284429 A1	21-06-2007 17-09-2008 07-05-2009 20-11-2008