Radester, A Novel Inhibitor of the Hsp90 Protein Folding Machinery

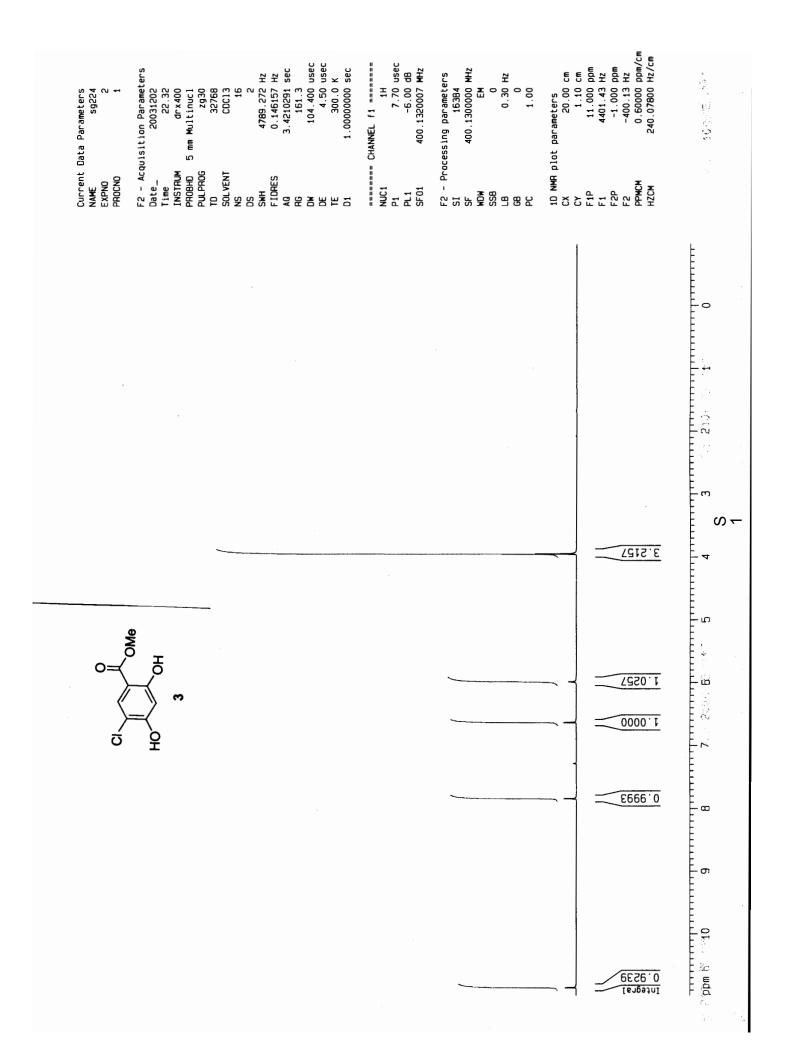
Gang Shen and Brian S. J. Blagg^{*}

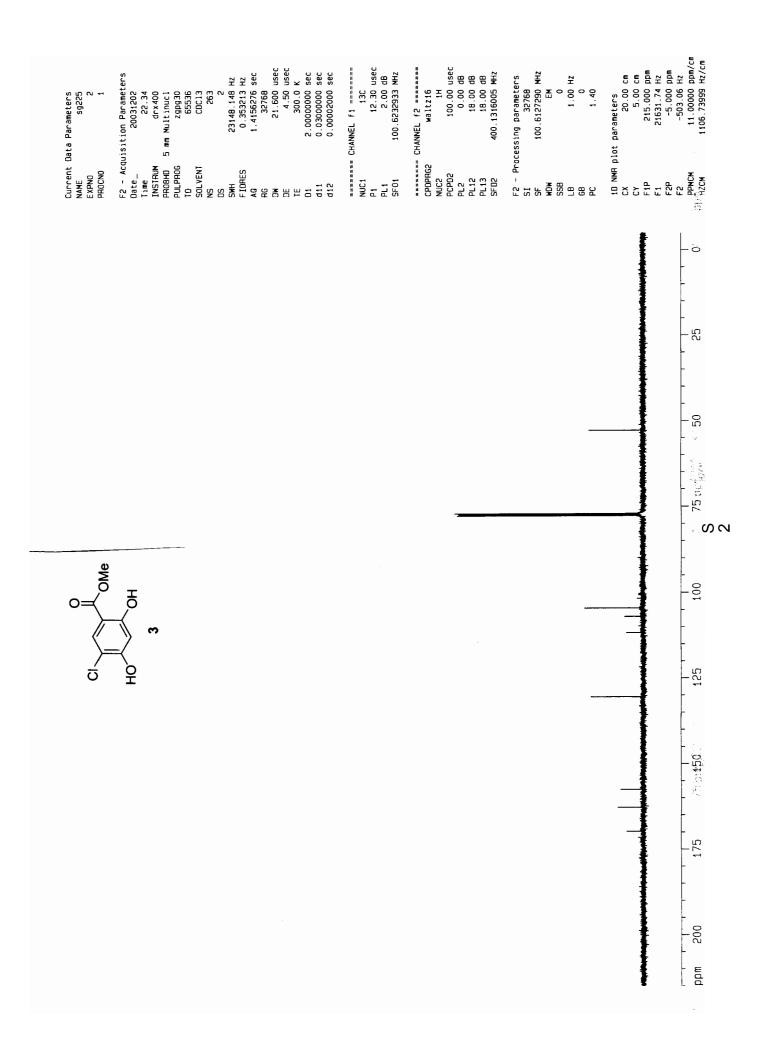
Department of Medicinal Chemistry and the Center for Protein Structure and Function 1251 Wescoe Hall Drive, Malott 4070, The University of Kansas, Lawrence, KS 66045-7563

bblagg@ku.edu

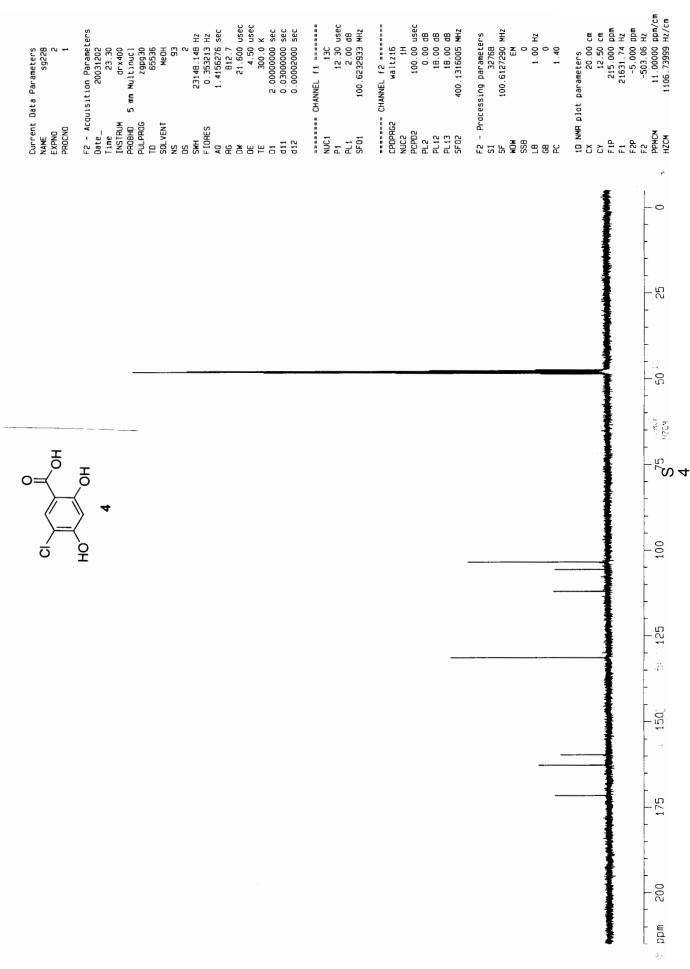
Supplementary Information

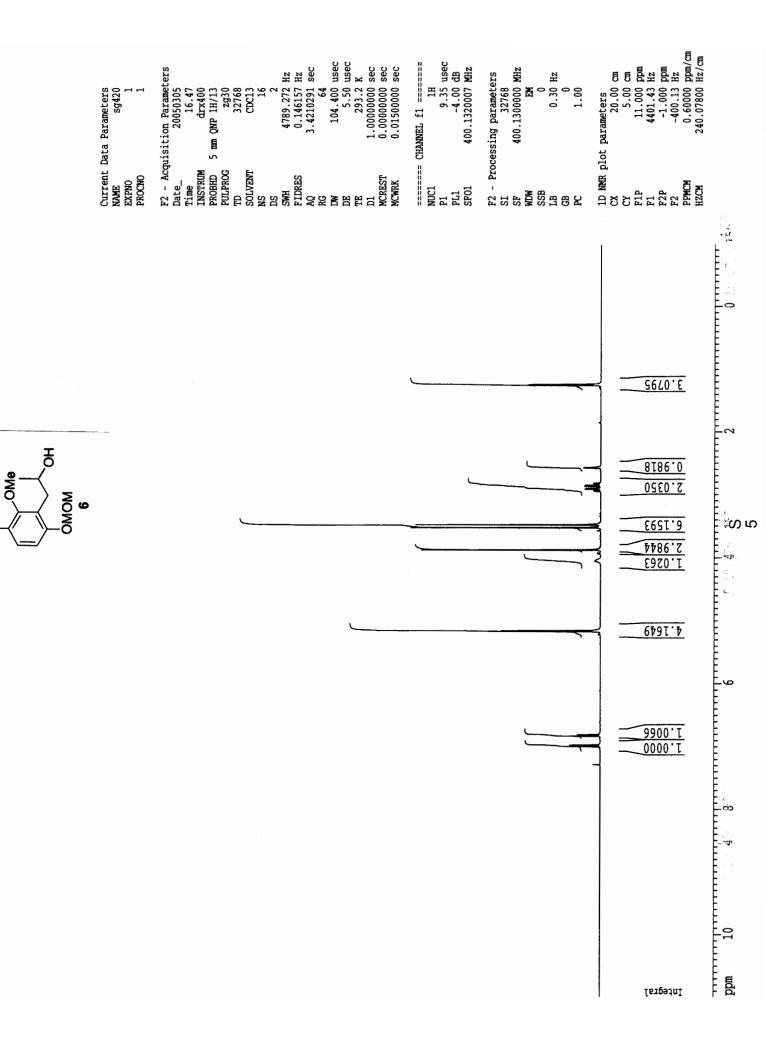
Spectra for Compounds



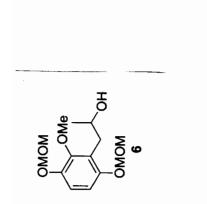


o <u>a</u>	л ш м.	All and a construct 1H NUC1 1H P1 7.70 usec PL1 7.70 usec PL3 7.00 dB SF01 400.1320007 MHz F2 Processing parameters SF 400.1300000 MHz SF 400.1300000 MHz SF 400.1300000 MHz BM 0 BB 0 CB 0 CB 0 CB 0 CC 1.00	JU NMH, plot parameters CX 20.00 cm CY 0.50 cm F1P 11.000 ppm F2P -1.000 ppm/cm F2P -0.013 Hz PPMCM 240.07800 Hz/cm HZCM 240.07800 Hz/cm
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			۵ ۲ 0573
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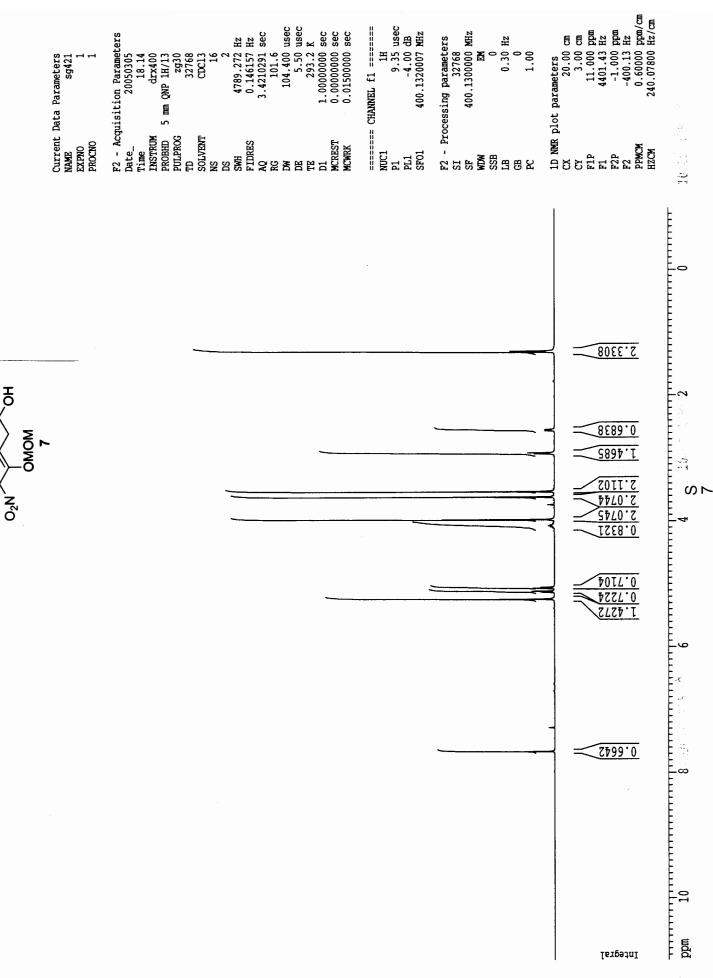




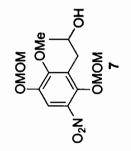
MOMO



ąć.



Current Data Parameters NAME sq421 EXENNO 2 PROCNO 1 F2 - Acquisition Parameters Date 2005005 Time 2005005 Time 2005000 PROBHD 5 mm QMP 1H/13 PRUBHD 5 mm QMP 1H/15 PRUCH 111.00000 Sec PRUBHD 5 mm QMP 1H/15 PRUCH 111.00000 PRUBHD 5 mm QMP 2000 PRUBHD 5 mm QMP 1H/15 PRUCH 111.00000 PRUBHD 5 mm QMP 2000 PRUCH 111.00000 PRUCH 20000 PRUCH 111.00000 PRUCH 20000 PRUCH 20000





Current Data Parameters NAME gs-572 EXPNO 1 PROCNO 1	F2 - Acquisition Parameters Date_ 20050307 Time 9.50 Time 2050307 Time 20050307 Time 205050307 Time 205050307 Date_ 20050307 Time 0.50 INSTRUM draveo TD 32768 SOLVENT 2930 NS 16 DS 2 SWH 4789.272 NS 0.146157 AQ 3.4210291 AQ 3.4210291 AQ 3.4210291 CB 3.4210291 CB 3.4210291	1: UEST 0. URK 0. 0. CHANNE	F2 - Processing parameters SI 32768 SF 400.1300000 MHz WDW EM 20 SSB 0 LB 0.30 Hz GB 0 PC 1.00	ID NMR plot parameters CX 20.00 cm CY 7.00 cm F1P 11.000 ppm F1 4401.43 Hz F2P -11.000 ppm F2P -400.13 Hz F2P 0.60000 ppm/cm HZCM 240.07800 Hz/cm	
				<u>E\$EE.E</u>	0
	80			<u> <i>FLIT.T</i></u> <i>FLIT.T FLIT.T FLIT.T</i>	≪.4.5 8.4 9
				<u>2.0886</u>	
				<u>1820.1</u>	8
					10
				Integral	und

 ID NMR plot parameters

 CX
 20.00 cm

 CY
 20.00 cm

 F1P
 215.000 ppm

 F2P
 -50.00 ppm

 F2P
 -500.00 ppm

 F2P
 -501.06 ppm

 F2M
 11.00000 ppm/cm
 ====== CHANNEL f2 ======= CPDPRG2 waltz16 NUC2 11 PCPD2 100.00 usec PL2 -4.00 dB PL12 16.58 dB PL13 17.00 dB SF02 400.1316005 MHz 23148.148 Hz 0.353213 Hz 1.4156276 sec 32768 usec 21.600 usec 23.5.50 usec 0.15000001 sec 0.015000000 sec 0.00000000 sec 0.015000000 sec 0.015000000 sec ====== CHANNEL f1 ======== NUC1 13C P1 11.25 usec PL1 2.00 dB SP01 100.6232933 MHz F2 - Processing parameters SI 65536 SF 100.6127290 MHZ MDW EM SSB 0 LB 1.00 Hz GB 0 FC 1.40
 F2
 - Acquisition Parameters

 Date______
 20050307

 Time
 20050307

 Time
 20050307

 Time
 12.53

 INSTRUM
 drx400

 PROBUD
 5 mm QNP 1H/13

 PULPROS
 299930

 TD
 05536

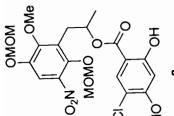
 SOLVENT
 05536

 NS
 1928

 NS
 1928

 NS
 23148.148

 FIDRES
 0.332313
 Current Data Parameters NAME gs-572 EXPNO 3 PROCNO 1 d11 DELTA MCREST MCWRK S 2 2 2 2 2 2 2 0 0 НŎ C 25 MOMO ω 5 50 È ΰ S 75 10 100 ÷ 125 150 175 DDm 200



 ID NMR plot parameters

 CX
 20.00 cm

 CY
 5.00 cm

 F1P
 11.000 ppm

 F1
 4401.43 Hz

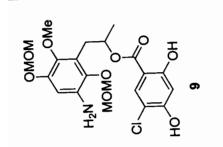
 F2P
 -1.000 ppm

 F2P
 -400.13 Hz

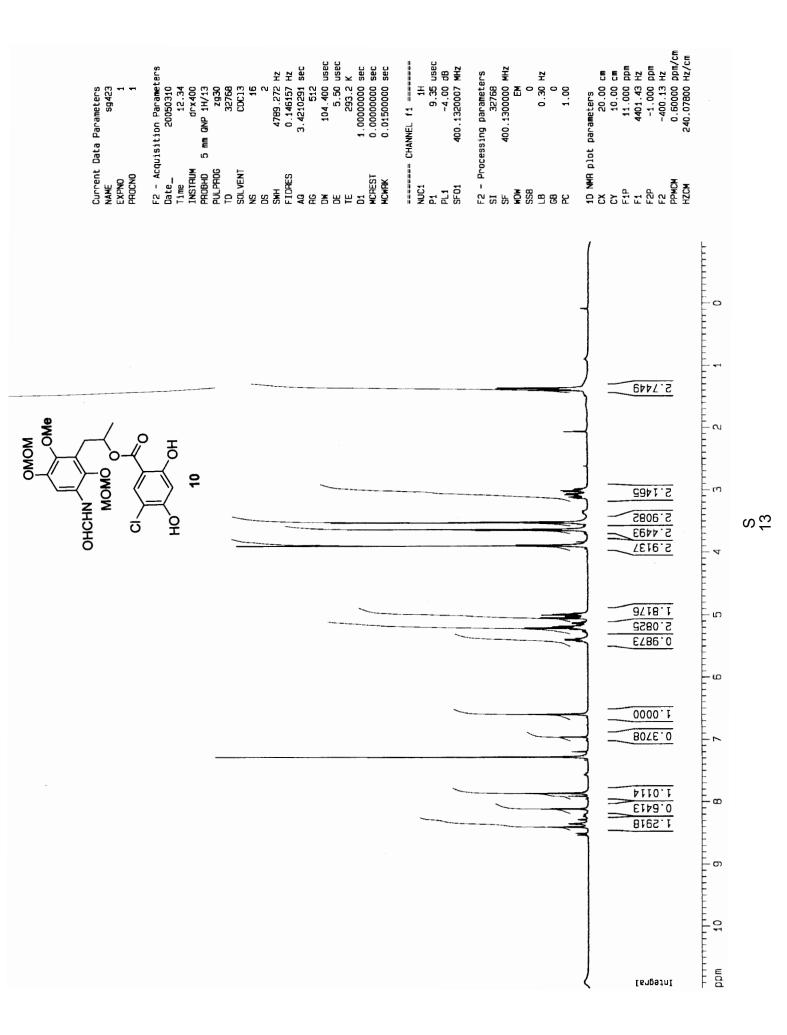
 PPMCM
 0.60000 ppm/cm

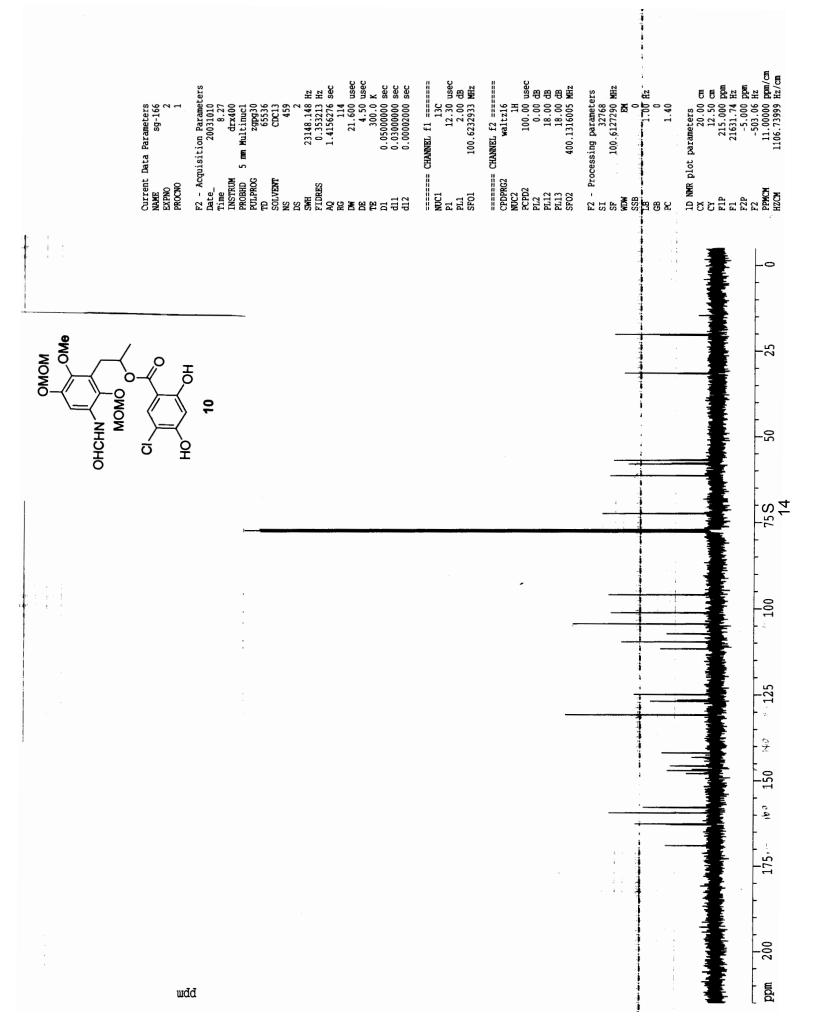
 HZCM
 240.07800 Hz/cm
 1 1 1 7.70 usec -6.00 dB 400.1320007 MHz 104.400 usec 4.50 usec F2 - Processing parameters S1 16384 SF 400.130000 MHz MDW EM 0 SSB 0.30 Hz G8 0.30 Hz G8 0.30 PZ 4789.272 Hz 0.146157 Hz 3.4210291 sec 101.6 300.0 K 1.00000000 sec CHANNEL f1 #22222 F2 - Acquisition Parameters Ħ 2930 32768 CDC13 16 2 16.48 drx400 sg-162 Current Data Parameters NAME sg-162 ru -5 mm Multinucl 20031007 INSTRUM PROBHD PUL PROG TD SOL VENT NS SWH PROCNO FIDRES Date_ EXPNO Time NUC1 P1 PL1 SF01 999 8 8 H H H 4134 B E286 8 OMe MOMO 0 د. 2 ترکیمی 4 میں 3 ع 1 میں میں 1 میں 1 میں 3 ع 0664 .E 9355 .E 9355 .E 9355 .E HO MOMO Ð s7 , N₂H ຮ່ È 5.1942 2.2521 1.1638 2.2029 1.0289 10.6916 mqq

	ers HAZ HAZ Sec USEC USEC SEC SEC	MH2 MH2 MH2 MH2 dB dB dB MH2	ers MHz Hz	cm cm ppm H2 H2 H2 H2 H2 Cm
.a Parameters sg-163 2	ittion Parameti 20031007 16.54 drx400 drx400 cgpggg 65536 CDC13 404 2.333213 1.4156276 1149.4 21.600 30.333213 1.4156276 0.333223 1.4156276 0.333223 0.333223 0.00000000 0.000000000	ANNEL f1 ==== 13C 12.30 2.00 2.00 2.00 12.30 12.30 12.00 13C 14 14 14 14 10 10 10 0.00 18.00 18.00 18.00	ssing paramete 32768 100.6127290 EM 1.00 1.40	t parameters 19.55 3.00 2055.00 2055.61 -10.036 11.00000 11.06.73999
Current Dat NAME EXPNO PROCNO	F2 - Acquis Date	CH NUC1 P1 P1 PL1 SF01 CPDPRG2 NUC2 NUC2 PL12 PL13 PL13 SF02 SF02	F2 - Proces SI MDW SSB SSB CB CB PC	10 NMA Plo CX CY F1P F2P F2P F2P F2P F2MCM H2CM









 1D NMR plot parameters

 CX
 20.00 cm

 CY
 12.00 cm

 F1P
 141.000 ppm

 F2P
 -4.000 ppm

 F2P
 -4.000 ppm

 F2P
 -4.000 ppm

 F2P
 -0.6000 ppm/cm

 HZCM
 240.07800 Hz/cm

 F2 - Acquisition Parameters

 Date _
 20050308

 Time
 9.43

 Time
 9.43

 INSTRIM
 0.7400

 PR00HD
 5 mm QNP 1H/13

 PR00HD
 5 mm QNP 1H/13

 PR0FND
 32768

 SOLVENT
 32768

 SOLVENT
 14/13

 PULPPDG
 32768

 SOLVENT
 1203769

 PULPPDG
 32768

 SOLVENT
 0.146157

 NS
 14/13

 NS
 14/13

 PULPPDG
 220013

 NS
 146157

 SOLVENT
 0.146157

 AQ
 3.4210291

 AG
 3.4210291

 PL
 0.146157

 AG
 3.4210291

 PL
 0.146157

 PL
 0.146157

 PL
 0.146157

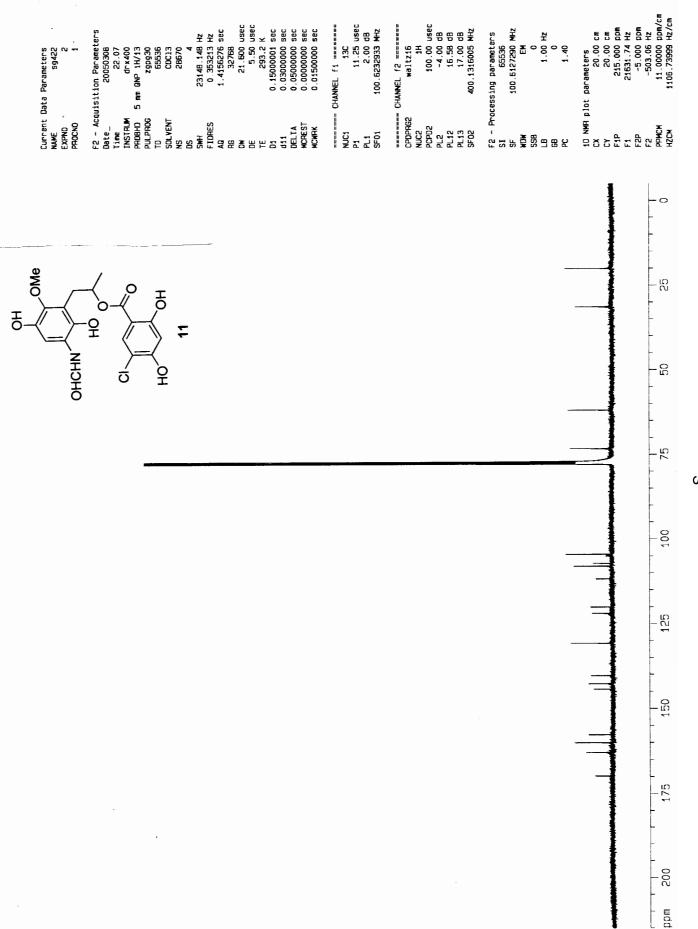
 PL
 0.146157

 PL
 0.146157

 PL
 0.01600000

 PL
 0.01500000

 PL
 0.0150 Current Data Parameters NAME gs-574 isopro hydro EXPNO 1 PROCNO 1 1H 9.35 usec -4.00 dB 400.1320007 MHz F2 - Processing parameters SI 32768 SF 400.1300000 MHz MDM EM SSB 0.30 Hz 68 0.30 Hz 68 0.30 Pz -----CHANNEL f1 ------NUC1 P1 PL1 SF01 0 T 3.5125 5.2929 OMe HO 4 S Ò ე ე 3.4409 Б P 7 OHCHN È Ω 1.3116 E-9 2890.1 1.0835 1.1782 2.0705 1.0831 1 . 0000 1 . 0000 шód



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 10 NMR plot parameters

 CX
 20.00 cm

 CY
 10.00 pm

 F1P
 11.000 ppm

 F2P
 -1.000 ppm

 F2P
 -400.13 Hz

 PPMCM
 0.60000 ppm/cm

 HZCM
 240.07800 Hz/cm
 F2 - Acquisition Parameters Date_______20031019 Time_______17.41 INSTRUM drx400 PHDBHD 5 mm Multinuc1 PULPHOG 2268 SOLVENT 2930 TD 2230 SOLVENT 20013 NS 16 OS 2 SMH 4789.272 Hz 181 104.400 usec 4.50 usec 300.0 K 1.0000000 sec 1H 7.70 usec -6.00 dB 400.1320007 MHz F2 - Processing parameters SI 16384 SF 400.1300000 MHZ MDW EM SSB 0 LB 0.30 HZ GB 0.30 HZ GB 7.00 2930 32768 22768 00013 16 2 4789.272 Hz 0.146157 Hz 0.146157 Hz 400.1300000 MHZ EM 0 0.30 Hz 1.00 ==== CHANNEL f1 ======= Current Data Parameters NAME sg-172 EXPNO 2 PRDCNO 1 FIORES 11111 NUC1 P1 PL1 SF01 8 원 8 원 분 5 3 2 1 0 3'3483 OMe 0 Ы Ю O: 1 . 0856 1 . 0525 OHCHN È ច s₽ 4 2.6634 1.2680 6960 · I د 8 ۲ 9087.0 LL96 0 1.1258 1.0200 10 1.0000 Integral шdd

20.00 cm 50.00 cm 215.00 ppm 21631.74 Hz -5.00 ppm -500 ppm -513.06 Hz 11.00000 ppm/cm 1106.73999 Hz/cm 23148.148 Hz 0.353213 Hz 1.4156276 sec 3768 21.600 usec 300.0 K 0.0500000 sec 0.0300000 sec 0.0002000 sec
 Emergence
 CHANNEL f1
 Emergence

 NUC1
 13C
 13C

 P1
 12.30 usec
 2.00 dB

 PL1
 100.6232933 MHz
 300 dB

 P2 - Acquisition Parameters

 Date
 20031019

 Time
 19.00

 Time
 19.00

 TNERCM
 drx400

 PROBHD
 5 mm Multinucl

 PULPROS
 297930

 TUL
 55536

 SULPAR
 7000

 DS
 23148.148

 HZ
 23148.148

 FIDRES
 0.353213
 P2- Processing parametersSI32768SP100.6127290 MHzNDMEMNDMEMSSB100.6127290 HzSSB0LB1.00 HzGB0PC1.40 THE CHANNEL F2 THE FILL 100.6127290 MHz Current Data Parameters NAME sg-173 EXFNO 2 PROCNO 1
 ID NMR plot parameters

 CX
 20.00

 CY
 50.00

 FIP
 215.000

 FIP
 21631.74

 F2P
 -55.000

 F2P
 -513.06

 F2PMCN
 11.00000

 H2CM
 1106.73999
 CPDPRG2 NUC2 PCPD2 PL2 PL12 PL13 SP02 Ю 22 OHCHN 오 $\overline{\mathbf{O}}$ 20 s ⁷⁵ 18 100 125 150 175 مرفعيل فمنعم كمر كمكسف أحاكم فكالكم كالكاكر والم 200 udd

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