

SUPPORTING INFORMATION

Supporting Information

Bioactive Compounds from *Crataegus pinnatifida* Bge. Leaves: Potential Health Benefits.

Rongrong Li ^{a,1}, Gaohui Shi ^{a,1}, Yue Liu ^a, Xinxin Lin ^b, Pinyi Gao ^c, Fang Wang ^b, Liqing Zhou ^a,
Lingzhi Li ^{a,*}

^a School of Traditional Chinese Materia Medica, Key Laboratory of Computational Chemistry-Based Natural Antitumor Drug Research & Development, Shenyang Pharmaceutical University, Shenyang, 110016, Liaoning Province, China

^b School of Functional Food and wine, Shenyang Pharmaceutical University, Wenhua Road 103, Shenyang 110016, Liaoning Province, China

^c College of Pharmaceutical and Biotechnology Engineering, Institute of Functional Molecules, Shenyang University of Chemical Technology, Shenyang 110142, China

¹ These authors contributed equally to this article.

* Corresponding author.

E-mail address: lilingzhijessie@163.com

SUPPORTING INFORMATION

Supporting Information

List of Contents

Figures

Fig S 1 :The scheme of isolation and activity of compounds from the leaves of <i>C. pinnatifida</i> Bge.....	5
Fig S 2 :Toxicity of compounds 1-21 at 25, 50, 100 μ M on the normal SH-SY5Y cells.	6
Fig S 3 :Assessment of apoptosis concentration induced by H ₂ O ₂ at different doses (100-900 μ M) for 24 h in SH-SY5Y cells.	7
Fig S 4 :Assessment of apoptosis concentration induced by ethanol at different doses (100-800 μ M) for 24 h in BRL-3A cell.....	7
Fig S 5 The HRESIMS spectrum of compound 1	8
Fig S 6 The ¹ H NMR spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 1	8
Fig S 7 The ¹³ C NMR spectrum (150 MHz, Methanol- <i>d</i> ₄) of compound 1	9
Fig S 8 The HSQC spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 1	9
Fig S 9 The HMBC spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 1	10
Fig S 10 The ROESY spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 1	10
Fig S 11 The CD spectrum (CDCl ₃) of compound 1'	11
Fig S 12 The HRESIMS spectrum of compound 2	11
Fig S 13 The ¹ H NMR spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 2	12
Fig S 14 The ¹³ C NMR spectrum (150 MHz, Methanol- <i>d</i> ₄) of compound 2	12
Fig S 15 The HMBC spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 2	13
Fig S 16 The CD spectrum (CDCl ₃) of compound 2'	13
Fig S 17 The HRESIMS spectrum of compound 3	14
Fig S 18 The ¹ H NMR spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 3	14
Fig S 19 The ¹³ C NMR spectrum (150 MHz, Methanol- <i>d</i> ₄) of compound 3	15
Fig S 20 The HSQC spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 3	15

SUPPORTING INFORMATION

Fig S 21 The HMBC spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 3	16
Fig S 22 The HRESIMS spectrum of compound 4	16
Fig S 23 The ¹ H NMR spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 4	17
Fig S 24 The ¹³ C NMR spectrum (150 MHz, Methanol- <i>d</i> ₄) of compound 4	17
Fig S 25 The HMBC spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 4	18
Fig S 26 The HRESIMS spectrum of compound 5	18
Fig S 27 The ¹ H NMR spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 5	19
Fig S 28 The ¹³ C NMR spectrum (150 MHz, Methanol- <i>d</i> ₄) of compound 5	19
Fig S 29 The HSQC spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 5	20
Fig S 30 The HMBC spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 5	20
Fig S 31 The NOESY spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 5	21
Fig S 32 The CD spectrum (MeOH) of compound 5'	21
Fig S 33 The HRESIMS spectrum of compound 6	22
Fig S 34 The ¹ H NMR spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 6	22
Fig S 35 The ¹³ C NMR spectrum (150 MHz, Methanol- <i>d</i> ₄) of compound 6	23
Fig S 36 The HSQC spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 6	23
Fig S 37 The HMBC spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 6	24
Fig S 38 The NOESY spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 6	24
Fig S 39 The CD spectrum (MeOH) of compound 6'	25
Fig S 40 The HRESIMS spectrum of compound 7	25
Fig S 41 The ¹ H NMR spectrum (600 MHz, DMSO- <i>d</i> ₆) of compound 7	26
Fig S 42 The ¹³ C NMR spectrum (150 MHz, DMSO- <i>d</i> ₆) of compound 7	26
Fig S 43 The HMBC spectrum (600 MHz, DMSO- <i>d</i> ₆) of compound 7	27
Fig S 44 The HRESIMS spectrum of compound 8	27
Fig S 45 The ¹ H NMR spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 8	28
Fig S 46 The ¹³ C NMR spectrum (150 MHz, Methanol- <i>d</i> ₄) of compound 8	28
Fig S 47 The HMBC spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 8	29
Fig S 48 The HRESIMS spectrum of compound 9	29
Fig S 49 The ¹ H NMR spectrum (600 MHz, Methanol- <i>d</i> ₄) of compound 9	30

SUPPORTING INFORMATION

Fig S 50 The ^{13}C NMR spectrum (150 MHz, Methanol- d_4) of compound 9 .	30
Fig S 51 The HMBC spectrum (600 MHz, Methanol- d_4) of compound 9 .	31
Fig S 52 The NOESY spectrum (600 MHz, Methanol- d_4) of compound 9 .	31
Fig S 53 The CD spectrum (MeOH) of compound 9 .	32
Fig S 54 The ^1H NMR spectrum (600 MHz, DMSO- d_6) of compound 10 .	32
Fig S 55 The ^{13}C NMR spectrum (150 MHz, DMSO- d_6) of compound 10 .	33
Fig S 56 The ^1H NMR spectrum (600 MHz, Methanol- d_4) of compound 11 .	33
Fig S 57 The ^{13}C NMR spectrum (150 MHz, Methanol- d_4) of compound 11 .	34
Fig S 58 The ^{13}C NMR spectrum (150 MHz, DMSO- d_6) of compound 12 .	34
Fig S 59 The ^{13}C NMR spectrum (150 MHz, DMSO- d_6) of compound 13 .	35
Fig S 60 The ^1H NMR spectrum (600 MHz, DMSO- d_6) of compound 14 .	35
Fig S 61 The ^{13}C NMR spectrum (150 MHz, DMSO- d_6) of compound 14 .	36
Fig S 62 The ^1H NMR spectrum (600 MHz, DMSO- d_6) of compound 15 .	36
Fig S 63 The ^{13}C NMR spectrum (150 MHz, DMSO- d_6) of compound 15 .	37
Fig S 64 The ^1H NMR spectrum (600 MHz, DMSO- d_6) of compound 16 .	37
Fig S 65 The ^{13}C NMR spectrum (150 MHz, DMSO- d_6) of compound 16 .	38
Fig S 66 The ^1H NMR spectrum (600 MHz, DMSO- d_6) of compound 17 .	38
Fig S 67 The ^{13}C NMR spectrum (150 MHz, DMSO- d_6) of compound 17 .	39
Fig S 68 The ^1H NMR spectrum (600 MHz, DMSO- d_6) of compound 18 .	39
Fig S 69 The ^{13}C NMR spectrum (150 MHz, DMSO- d_6) of compound 18 .	40
Fig S 70 The ^1H NMR spectrum (600 MHz, DMSO- d_6) of compound 19 .	40
Fig S 71 The ^{13}C NMR spectrum (150 MHz, DMSO- d_6) of compound 19 .	41
Fig S 72 The ^{13}C NMR spectrum (150 MHz, DMSO- d_6) of compound 20 .	41
Fig S 73 The ^{13}C NMR spectrum (150 MHz, DMSO- d_6) of compound 21 .	42

Tables

Table S 1 ^1H (600 MHz) and ^{13}C (150 MHz) data of compounds 1 and 2 (Methanol- d_4).	43
Table S 2 ^1H (600 MHz) and ^{13}C (150 MHz) data of compounds 3 and 4 (Methanol- d_4).	44
Table S 3. [α] data of compounds 3 and 4 at 20 °C with 589 nm laser (methanol as the solvent) and the	

SUPPORTING INFORMATION

calculated specific rotation of their aglycone in four absolute configurations.....	45
Table S 4 ^1H (600 MHz) and ^{13}C (150 MHz) data of compounds 5 and 6 (Methanol- d_4).....	46
Table S 5 ^1H (600 MHz) and ^{13}C (150 MHz) data of compound 7 (DMSO- d_6).....	47
Table S 6 ^1H (600 MHz) and ^{13}C (150 MHz) data of compound 8 (Methanol- d_4).....	48
Table S 7 ^1H (600 MHz) and ^{13}C (150 MHz) data of compound 9 (Methanol- d_4).....	49
Table S 8 ^1H (600 MHz) and ^{13}C -NMR (150 MHz) data of compounds 10 and 11	50
Table S 9 ^{13}C -NMR (150 MHz) data of compounds 12-15 (DMSO- d_6).....	52
Table S 10 ^{13}C -NMR (150 MHz) data of compounds 16-21 (DMSO- d_6).....	53
Table S 11 Antioxidant activity of separation fractions of the leaves of <i>Crataegus pinnatifida</i> Bge. <i>in vitro</i>	54
Table S 12 Antioxidant activity of compounds 1-21 <i>in vitro</i>	55
Table S 13 The protective effect of flavonoids 1-21 on SH-SY5Y cells at 25, 50, 100 μM	56
Table S 14 The protective effect of flavonoids 10, 12-21 on alcoholic hepatocyte injury at 25, 50, 100 μM	57

SUPPORTING INFORMATION

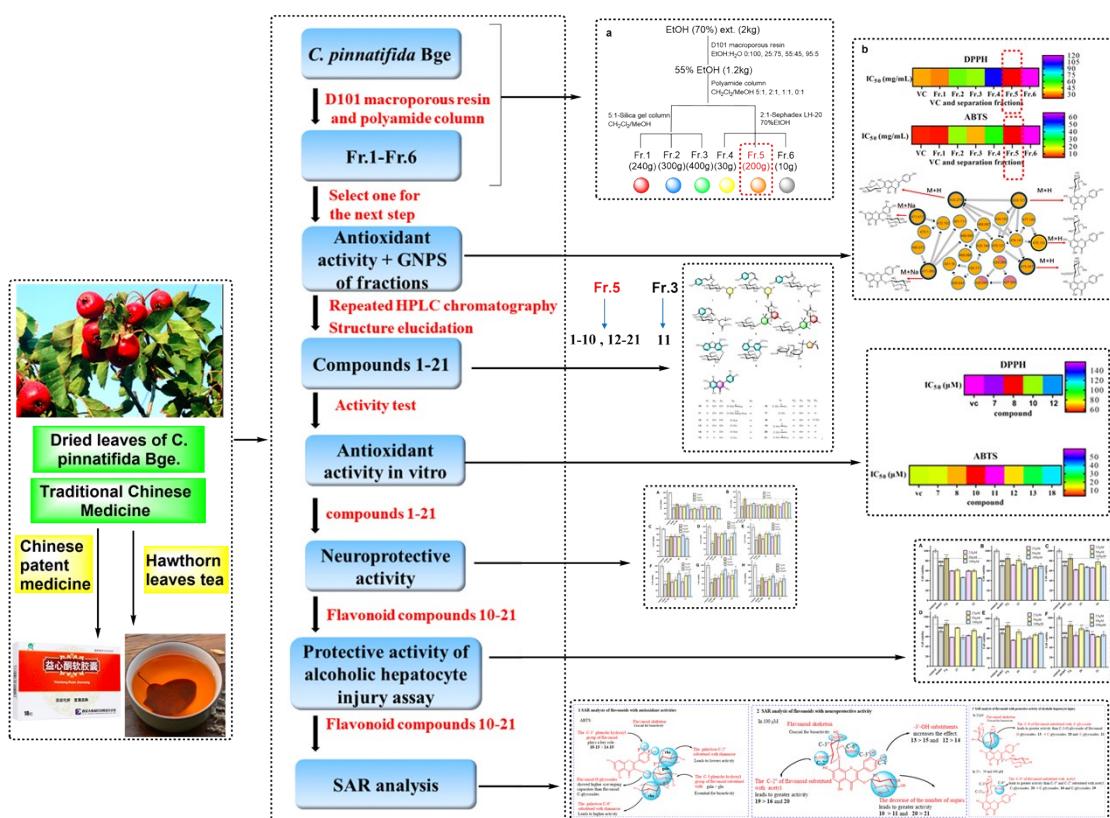


Fig S 1 :The scheme of isolation and activity of compounds from the leaves of *C. pinnatifida* Bge

SUPPORTING INFORMATION

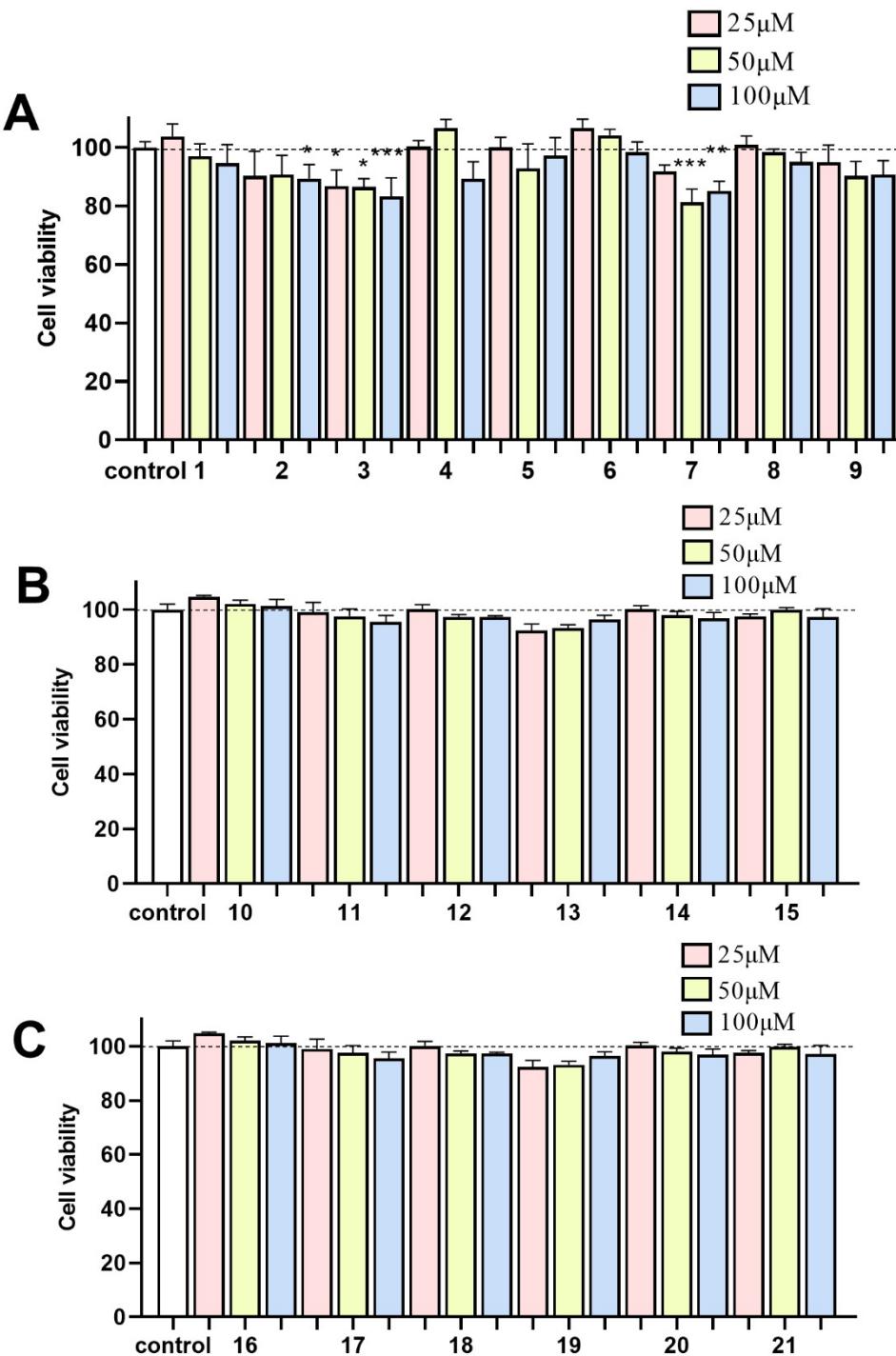


Fig S 2 :Toxicity of compounds 1-21 at 25, 50, 100 μ M on the normal SH-SY5Y cells.

(In this figure, "control" is the blank group. Values are presented as percentage differences from control and they are significantly different with $P>0.05$ for Student's t-test: * $P < 0.5$, ** $P < 0.01$, and *** $P < 0.001$. The bars represent mean \pm SD ($n=3$)).

SUPPORTING INFORMATION

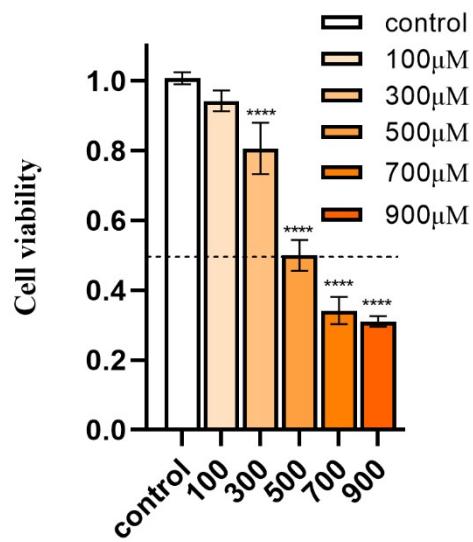


Fig S 3 :Assessment of apoptosis concentration induced by H₂O₂ at different doses (100-900 μ M) for 24 h in SH-SY5Y cells.

The value is the mean \pm SD of three replicates, **** $P < 0.0001$ vs control. The bars represent mean \pm SD ($n=3$).

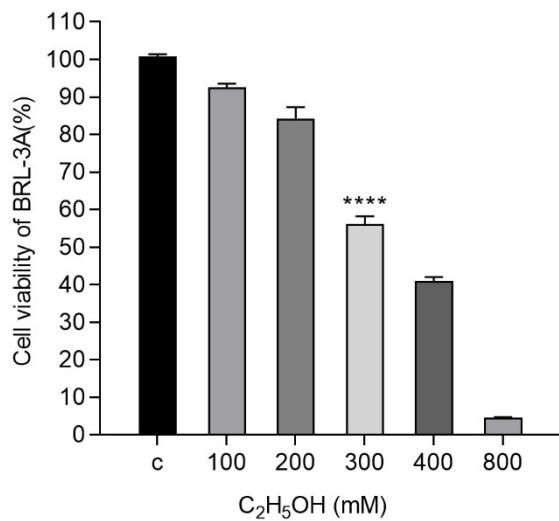


Fig S 4 :Assessment of apoptosis concentration induced by ethanol at different doses (100-800 μ M) for 24 h in BRL-3A cell.

Values were the mean \pm SD of three replicates, **** $P < 0.0001$ vs control group. The bars represent mean \pm SD ($n=3$).

SUPPORTING INFORMATION

Compound Spectrum SmartFormula Report

Analysis Info		Acquisition Date	12/27/2021 2:28:10 PM
Analysis Name	D:\Data\YIJI\2021\227\SY-35_8_1_3117.d	Operator	Demo User
Method	HPLC_MS_pos_without_column.m	Instrument	compact
Sample Name	SY-35		8255754.20225
Comment			

Acquisition Parameter					
Source Type	ESI	Ion Polarity	Positive	Set Nebulizer	1.8 Bar
Focus	Not active	Set Capillary	4500 V	Set Dry Heater	220 °C
Scan Begin	50 m/z	Set End Plate Offset	-500 V	Set Dry Gas	8.0 l/min
Scan End	1300 m/z	Set Charging Voltage	2000 V	Set Divert Valve	Waste
		Set Corona	0 nA	Set APCI Heater	0 °C

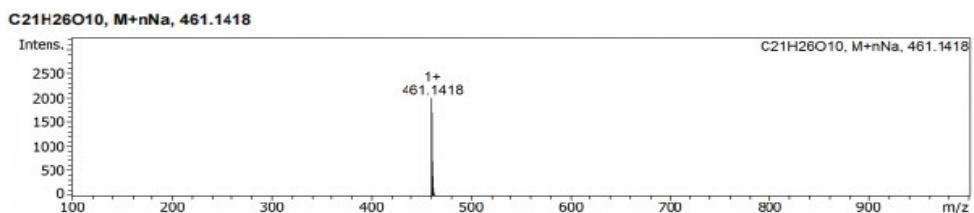
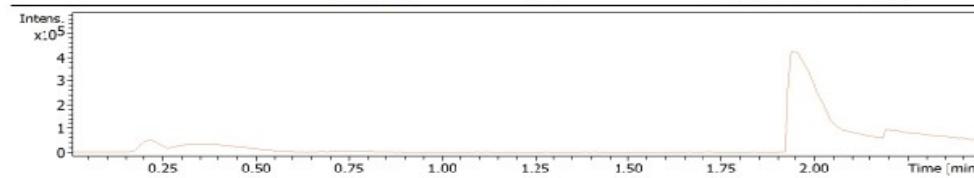


Fig S 5 The HRESIMS spectrum of compound 1.

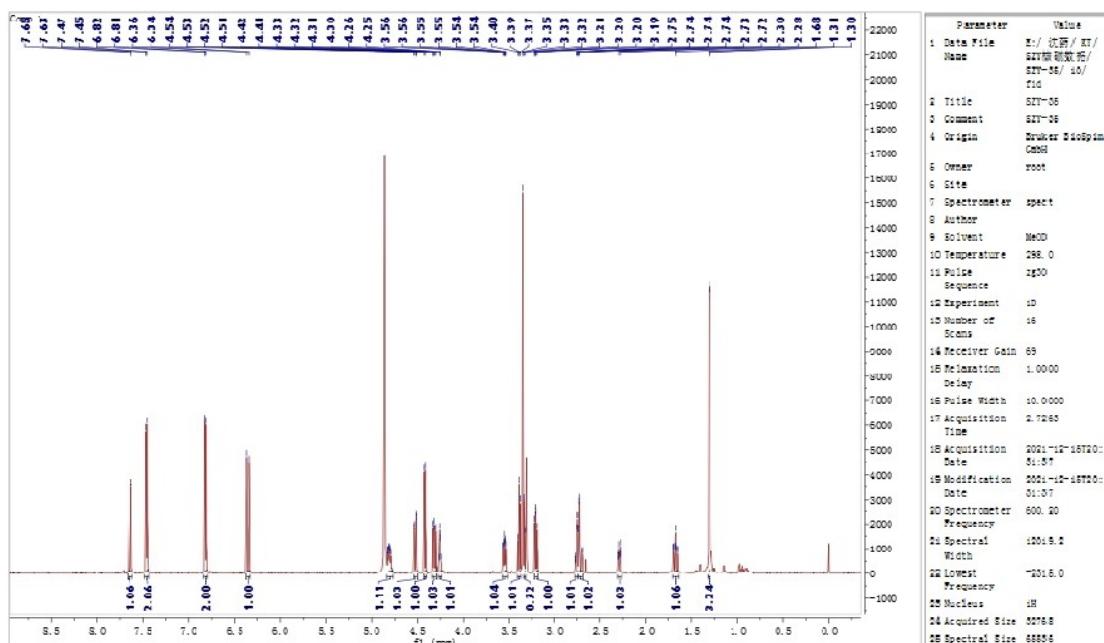


Fig S 6 The ¹H NMR spectrum (600 MHz, Methanol-d₄) of compound 1.

SUPPORTING INFORMATION

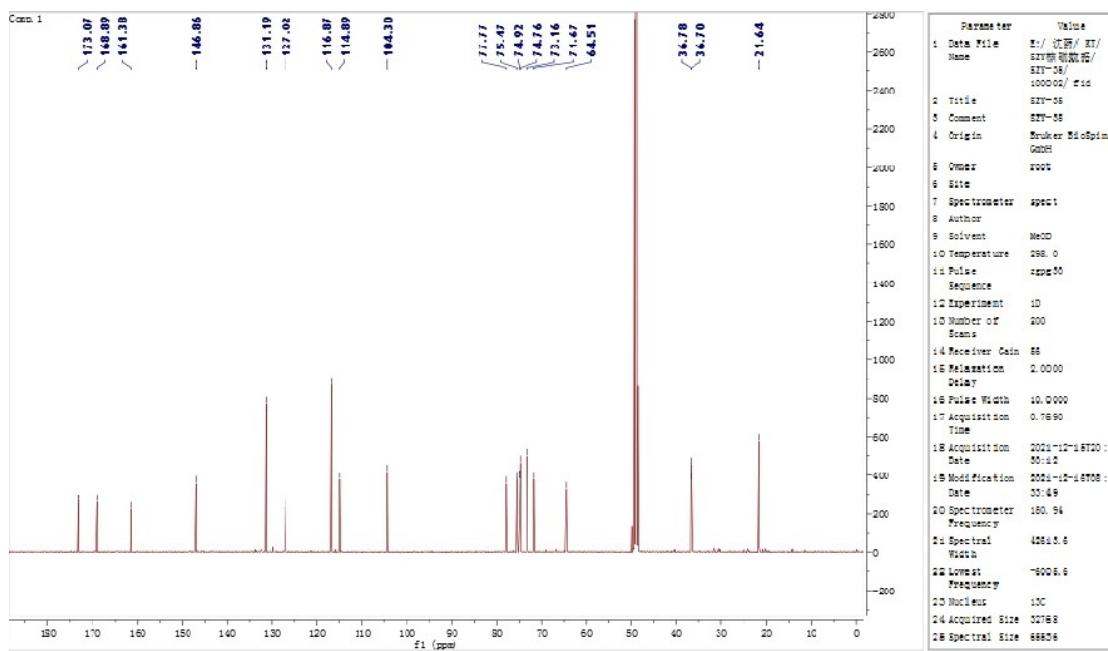


Fig S 7 The ^{13}C NMR spectrum (150 MHz, Methanol- d_4) of compound 1.

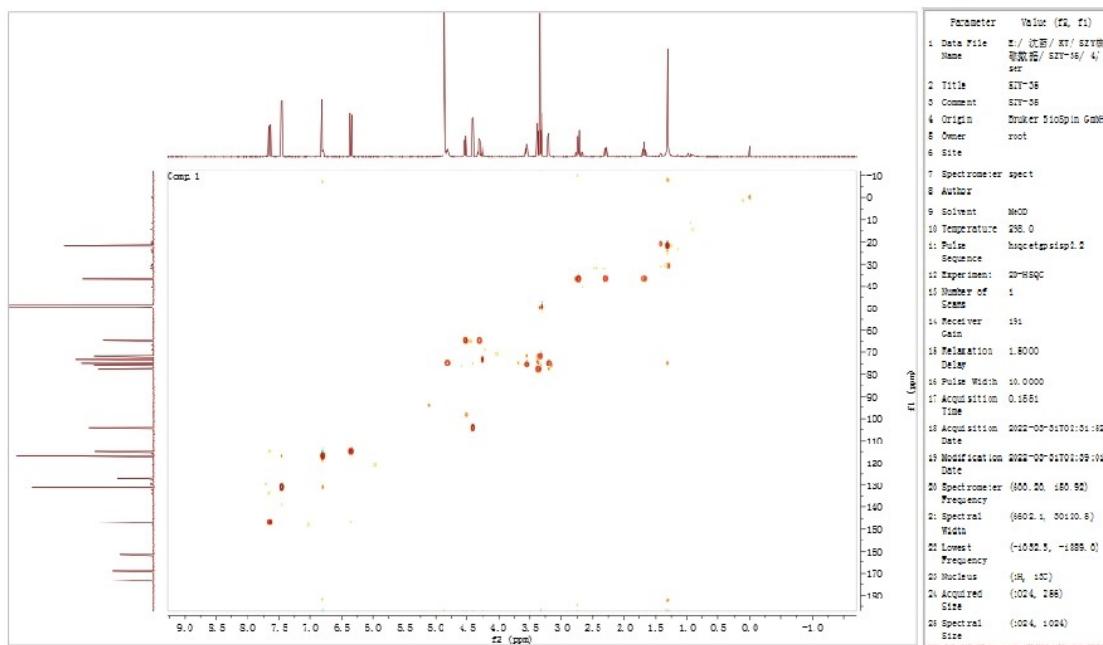


Fig S 8 The HSQC spectrum (600 MHz, Methanol- d_4) of compound 1.

SUPPORTING INFORMATION

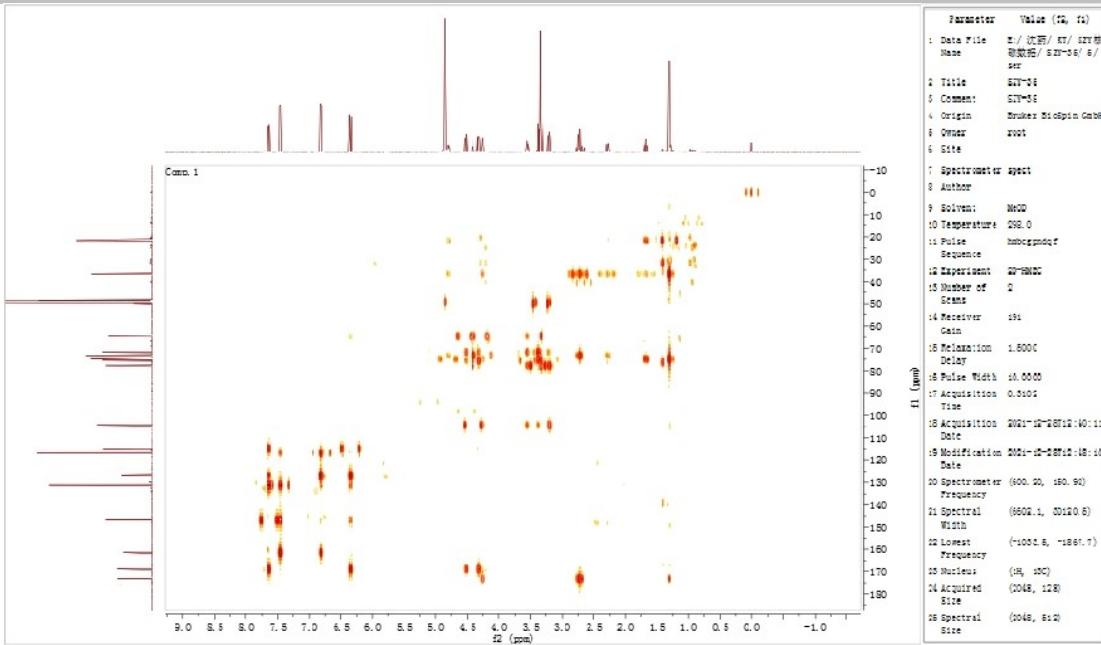


Fig S 9 The HMBC spectrum (600 MHz, Methanol-*d*₄) of compound 1.

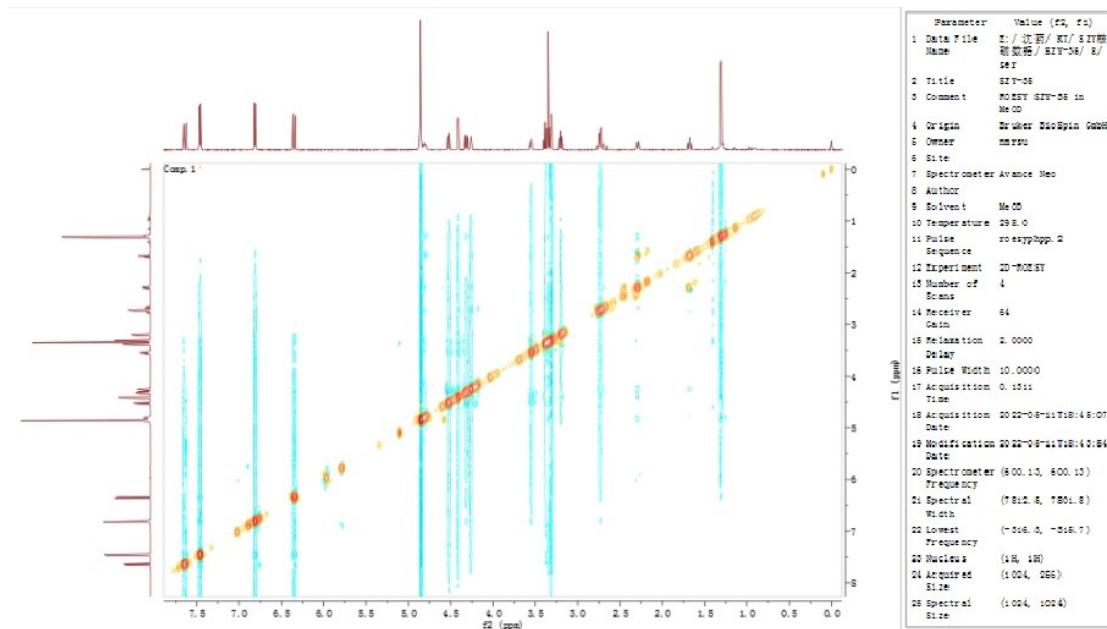


Fig S 10 The ROESY spectrum (600 MHz, Methanol-*d*₄) of compound 1.

SUPPORTING INFORMATION

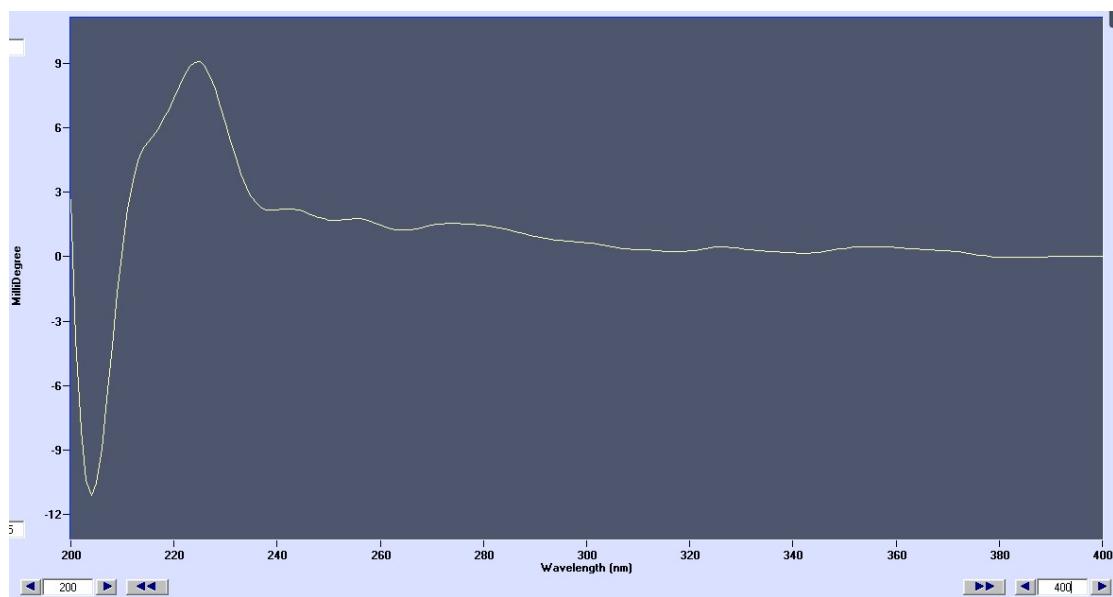


Fig S 11 The CD spectrum (CDCl_3) of compound **1'**.

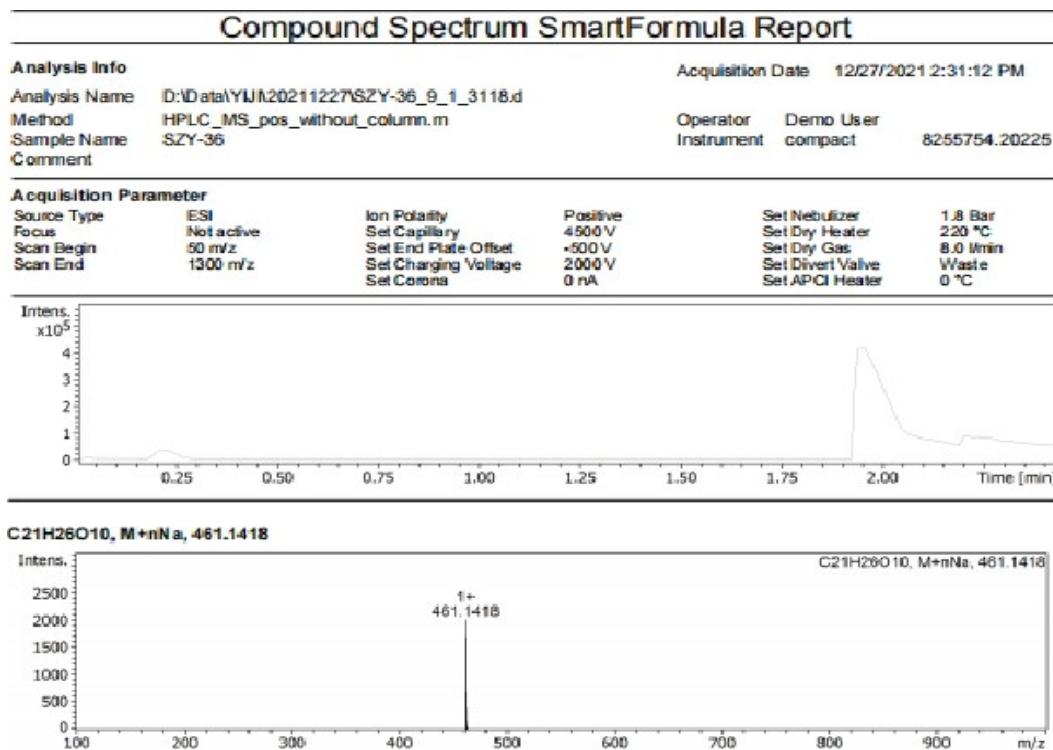


Fig S 12 The HRESIMS spectrum of compound **2**.

SUPPORTING INFORMATION

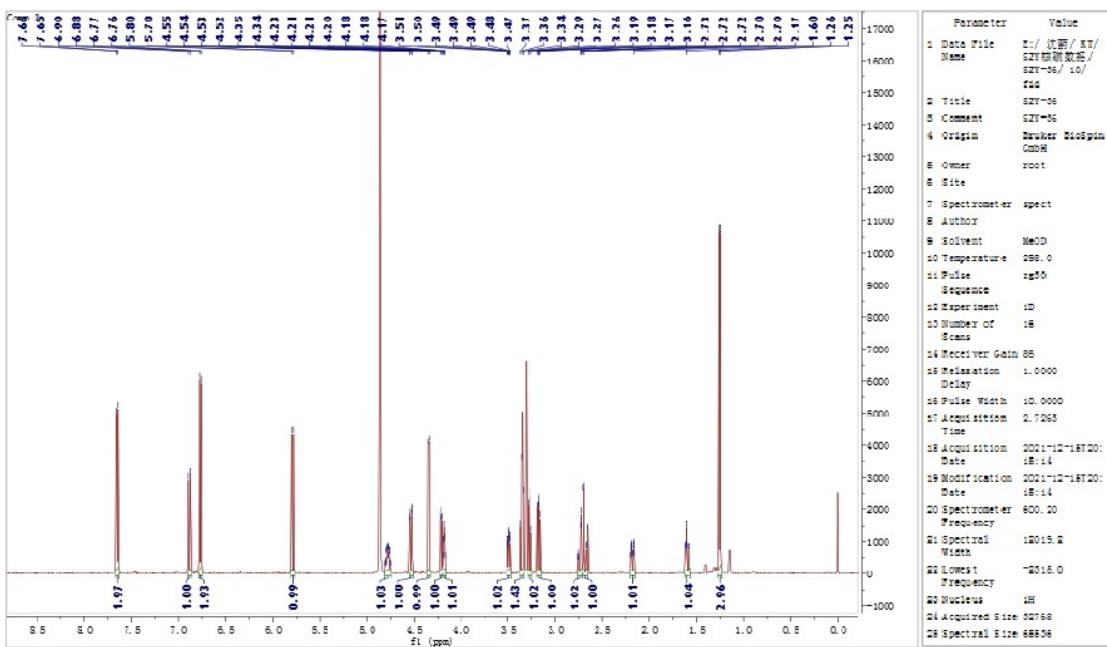


Fig S 13 The ^1H NMR spectrum (600 MHz, Methanol- d_4) of compound 2.

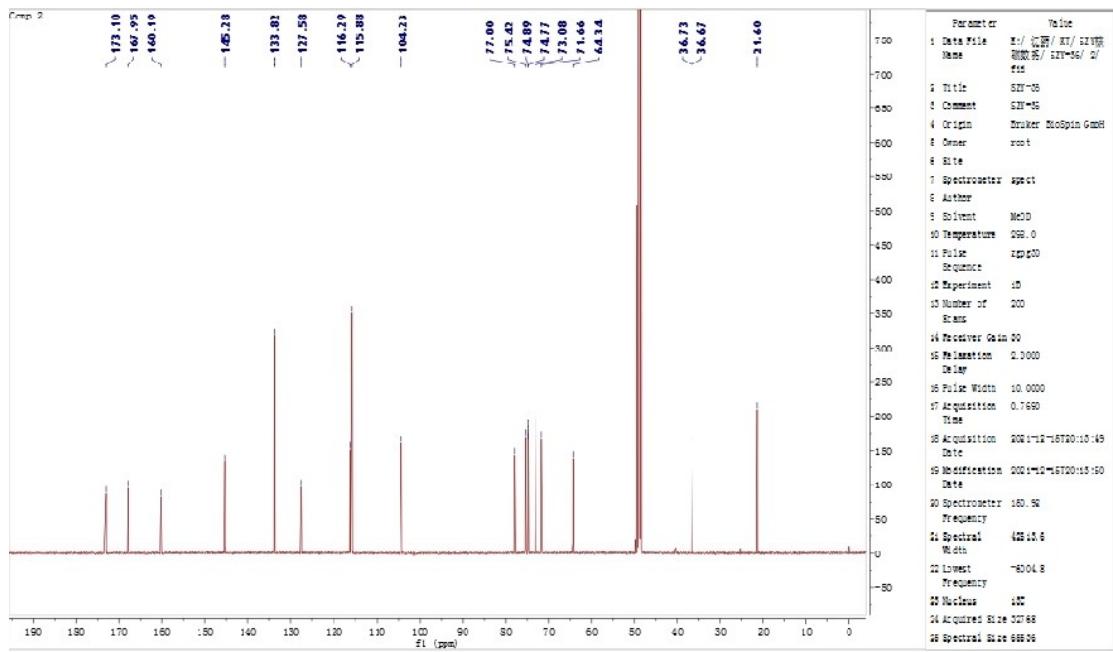


Fig S 14 The ^{13}C NMR spectrum (150 MHz, Methanol- d_4) of compound 2.

SUPPORTING INFORMATION

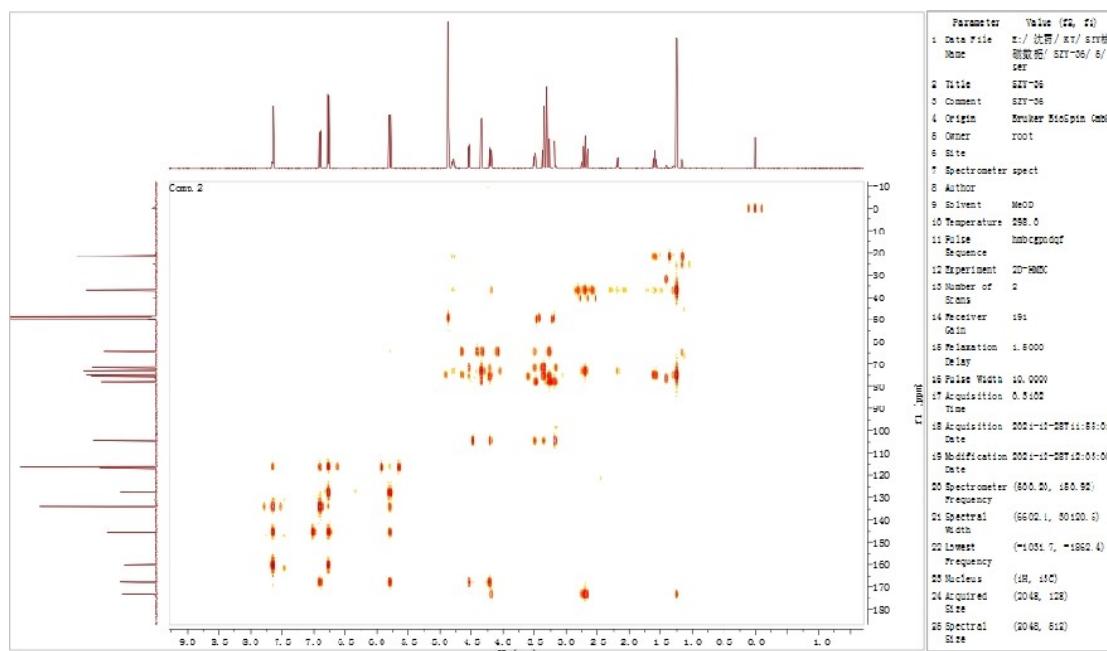


Fig S 15 The HMBC spectrum (600 MHz, Methanol- d_4) of compound 2.

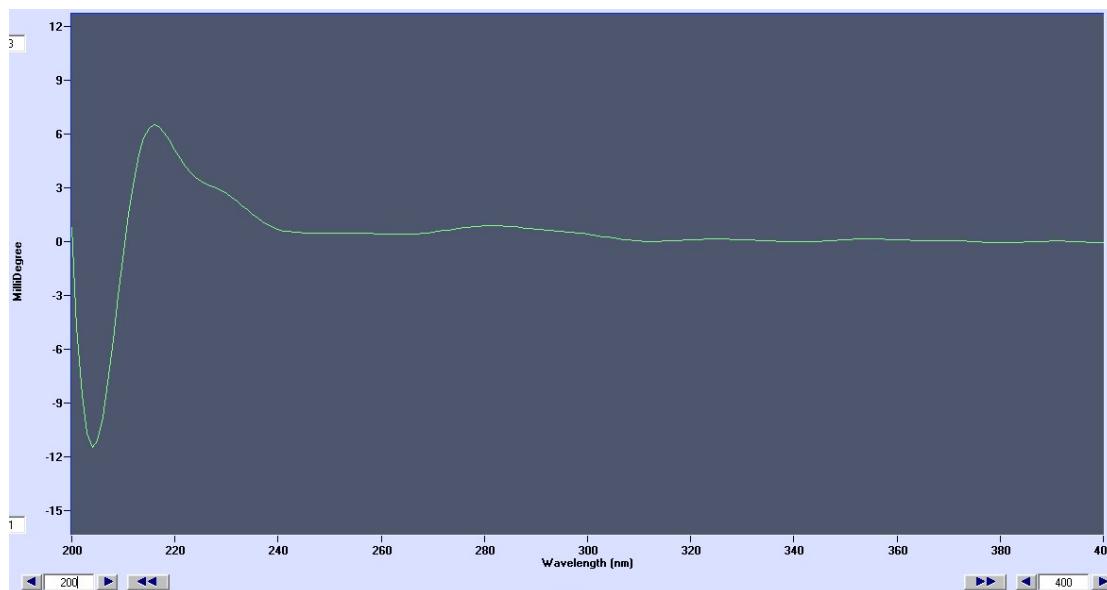
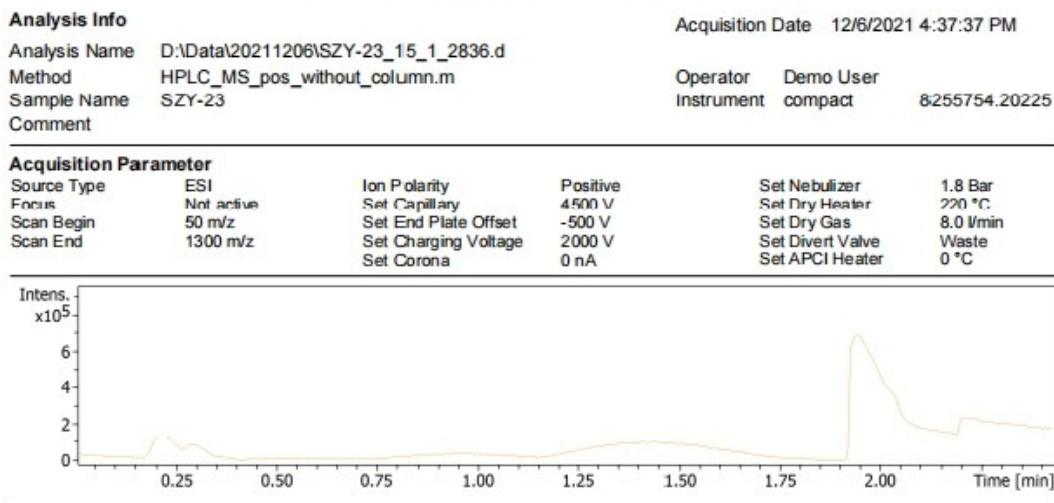


Fig S 16 The CD spectrum (CDCl_3) of compound 2'.

SUPPORTING INFORMATION

Compound Spectrum SmartFormula Report



C22H30O11, M+nNa, 493.1680

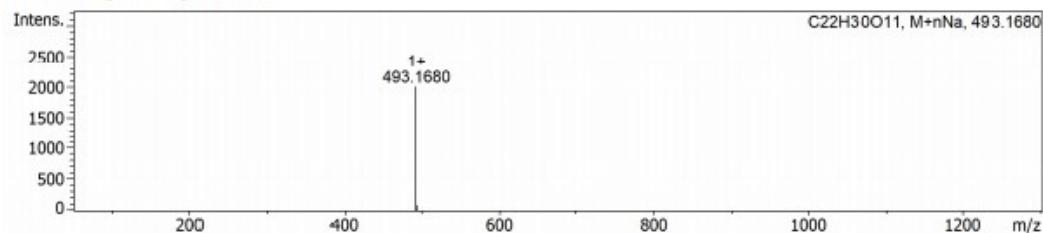


Fig S 17 The HRESIMS spectrum of compound 3.

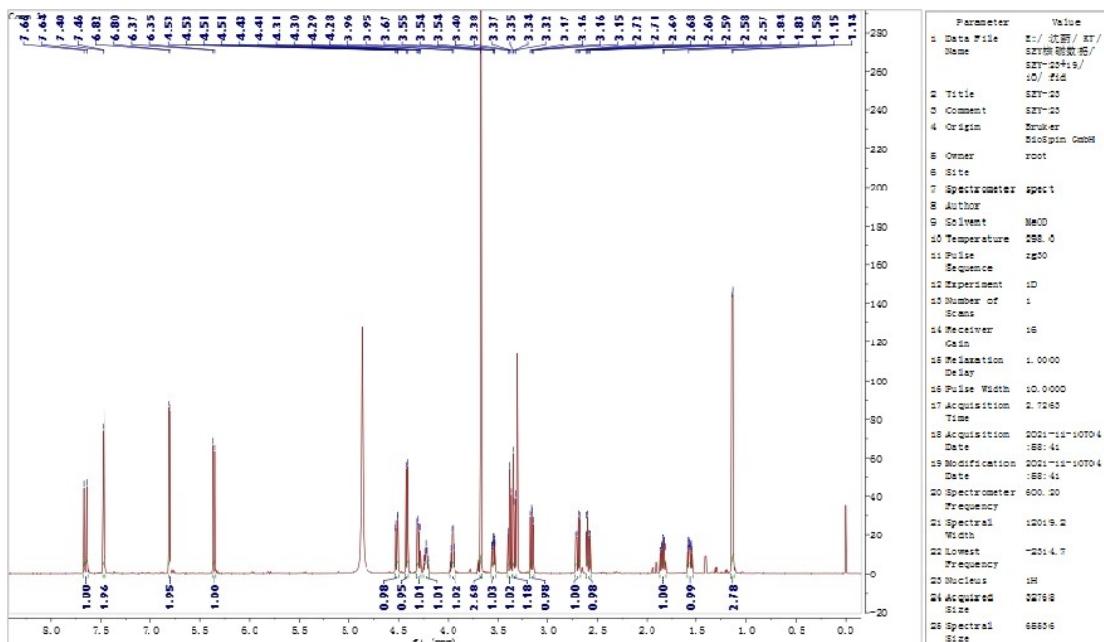


Fig S 18 The ¹H NMR spectrum (600 MHz, Methanol-d₄) of compound 3.

SUPPORTING INFORMATION

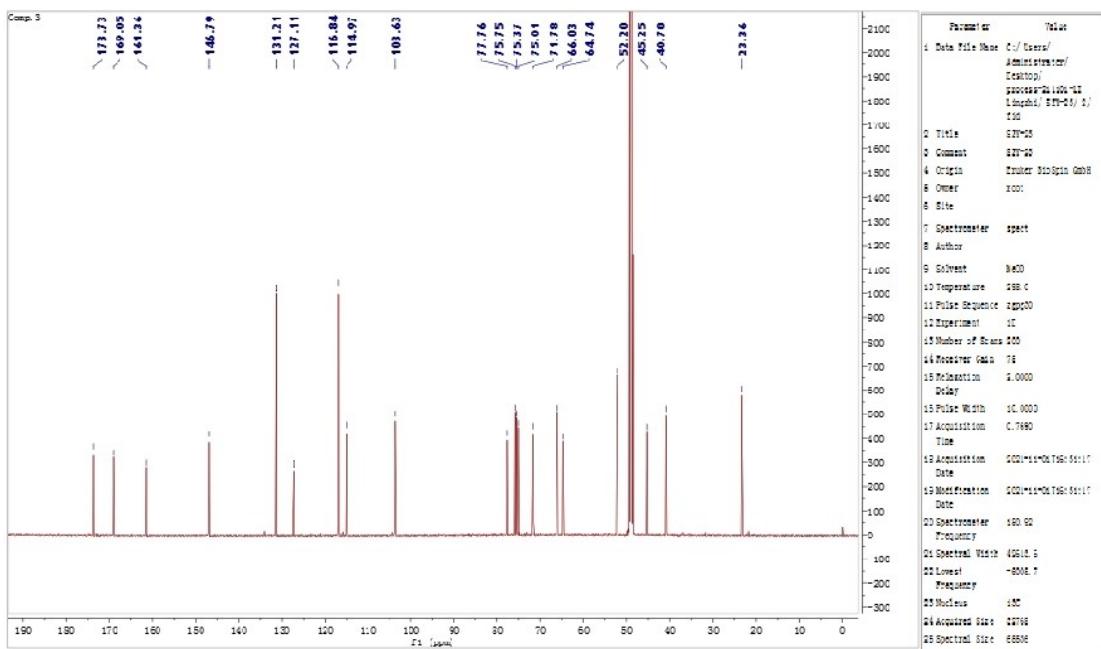


Fig S 19 The ^{13}C NMR spectrum (150 MHz, Methanol- d_4) of compound 3.

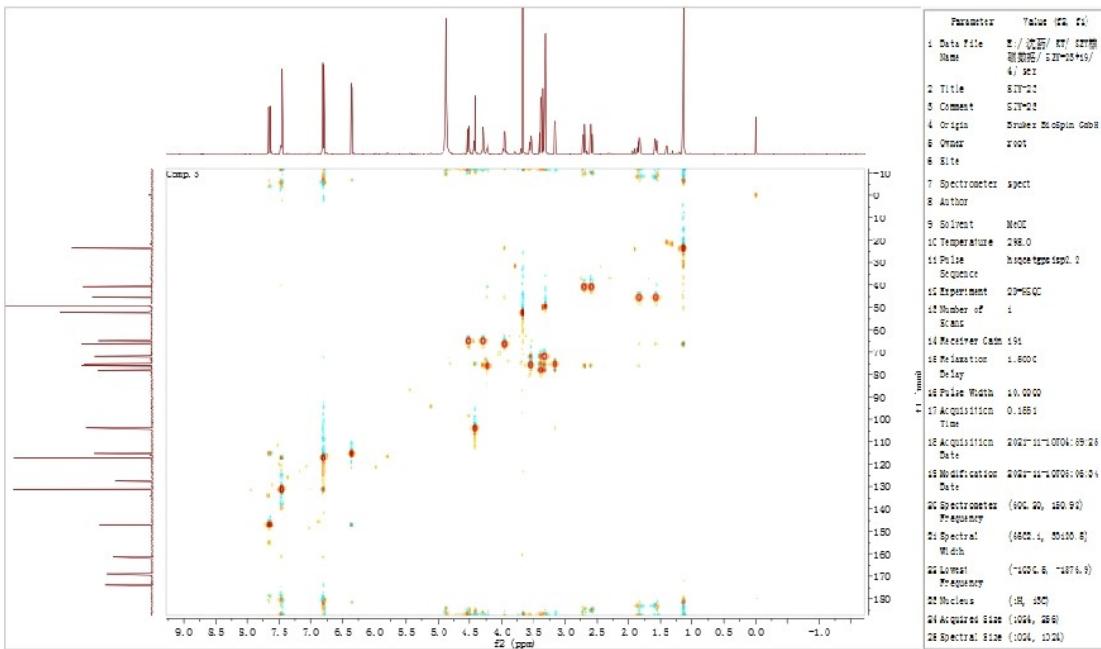


Fig S 20 The HSQC spectrum (600 MHz, Methanol- d_4) of compound 3.

SUPPORTING INFORMATION

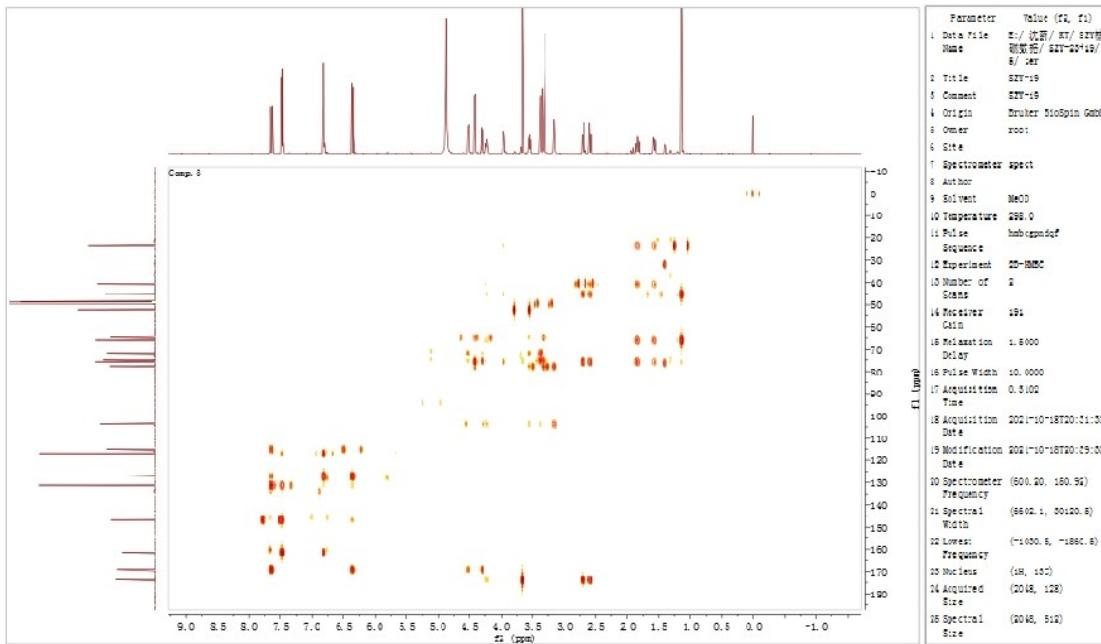


Fig S 21 The HMBC spectrum (600 MHz, Methanol-*d*₄) of compound 3.

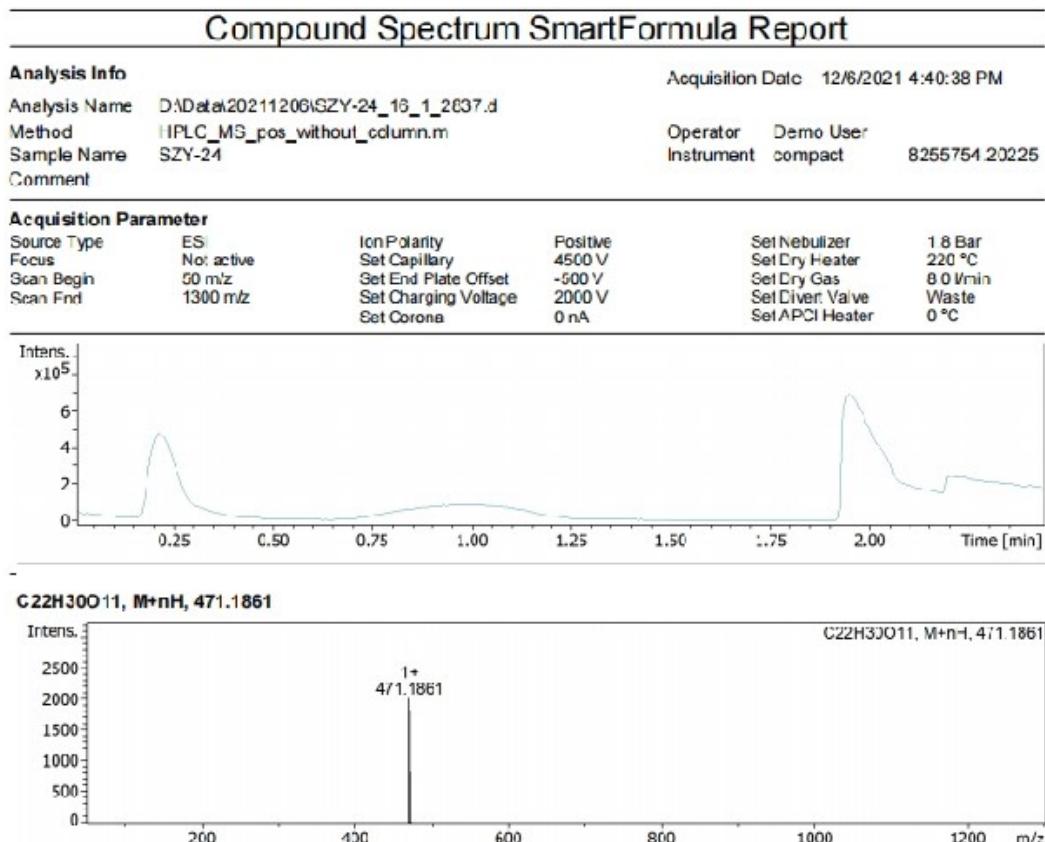


Fig S 22 The HRESIMS spectrum of compound 4.

SUPPORTING INFORMATION

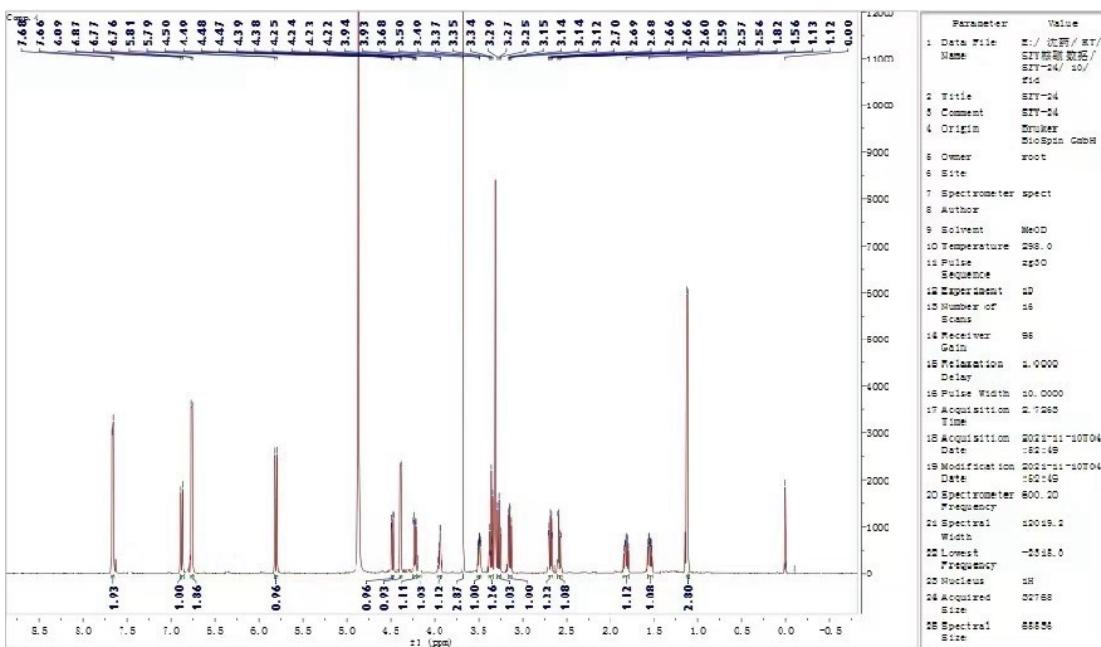


Fig S 23 The ¹H NMR spectrum (600 MHz, Methanol-*d*₄) of compound 4.

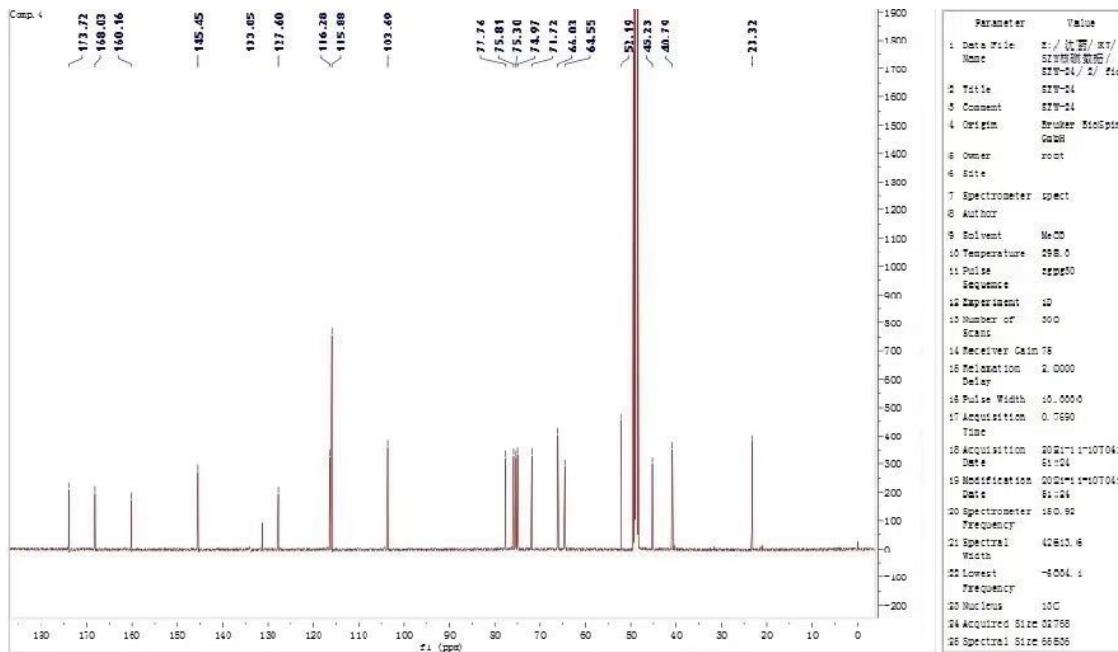


Fig S 24 The ¹³C NMR spectrum (150 MHz, Methanol-*d*₄) of compound 4.

SUPPORTING INFORMATION

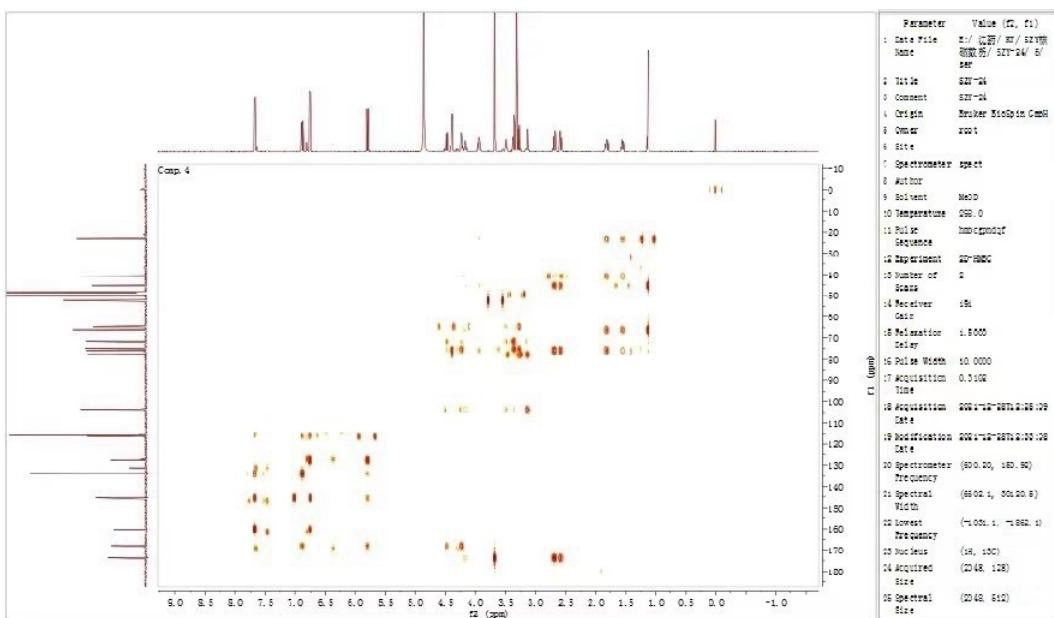


Fig S 25 The HMBC spectrum (600 MHz, Methanol-*d*₄) of compound 4.

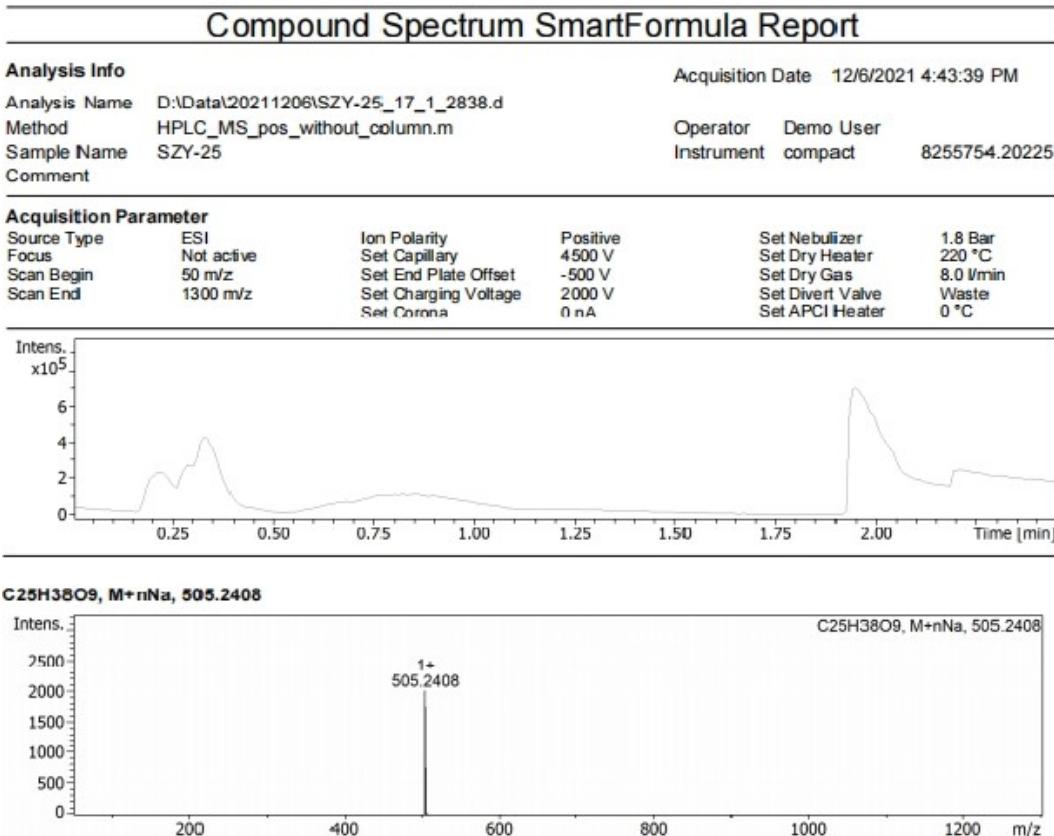


Fig S 26 The HRESIMS spectrum of compound 5.

SUPPORTING INFORMATION

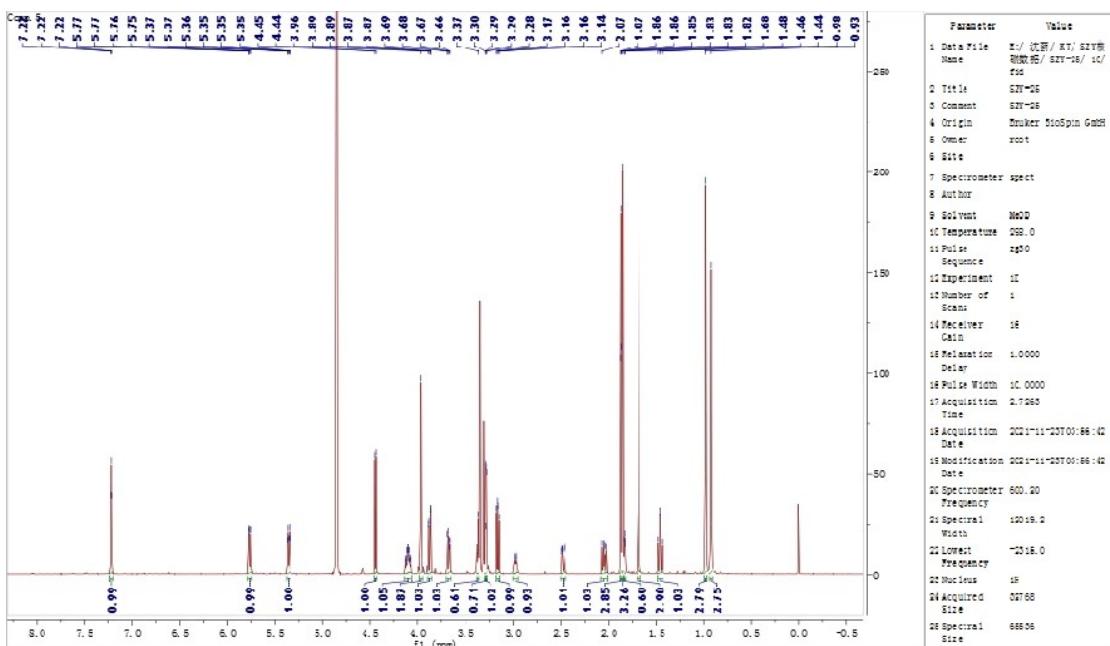


Fig S 27 The ^1H NMR spectrum (600 MHz, Methanol- d_4) of compound 5.

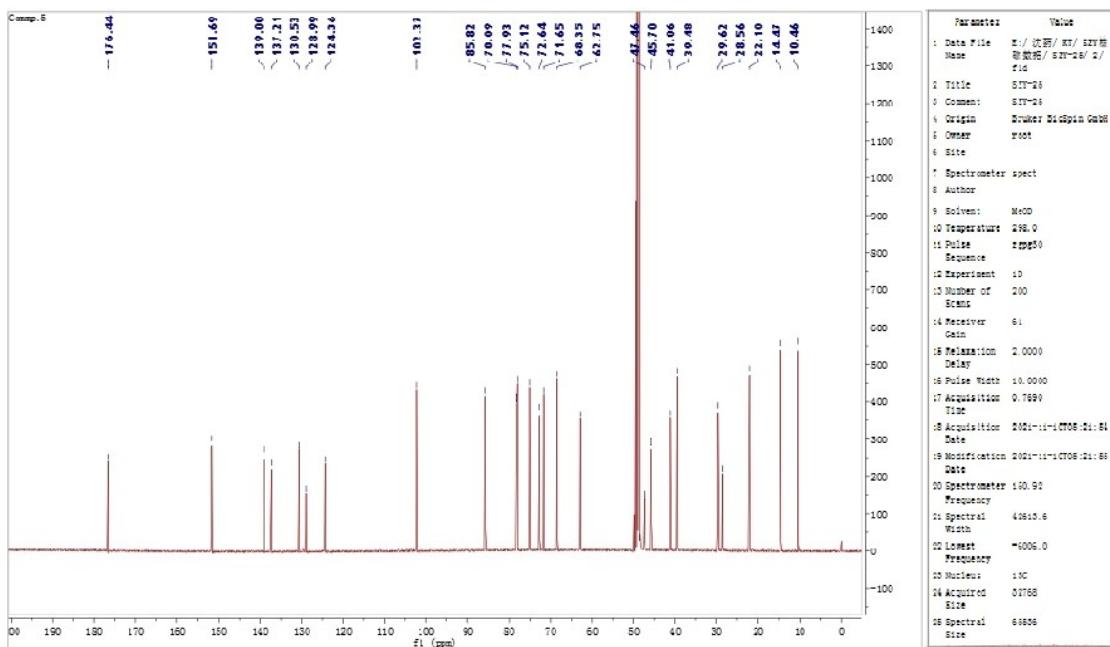


Fig S 28 The ^{13}C NMR spectrum (150 MHz, Methanol- d_4) of compound 5.

SUPPORTING INFORMATION

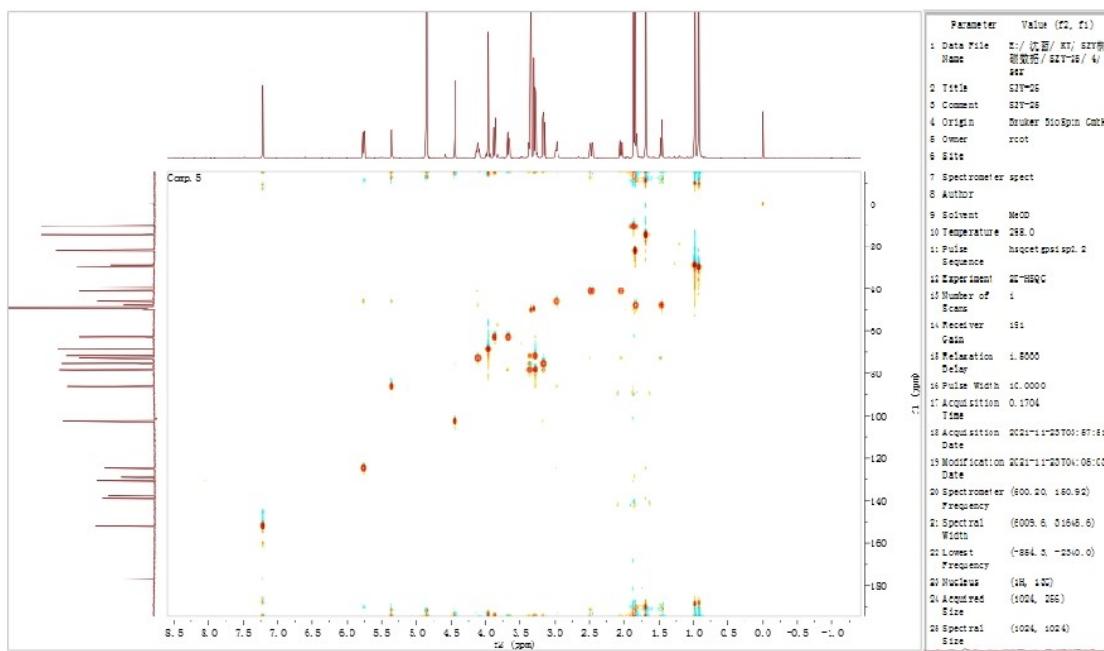


Fig S 29 The HSQC spectrum (600 MHz, Methanol-*d*₄) of compound 5.

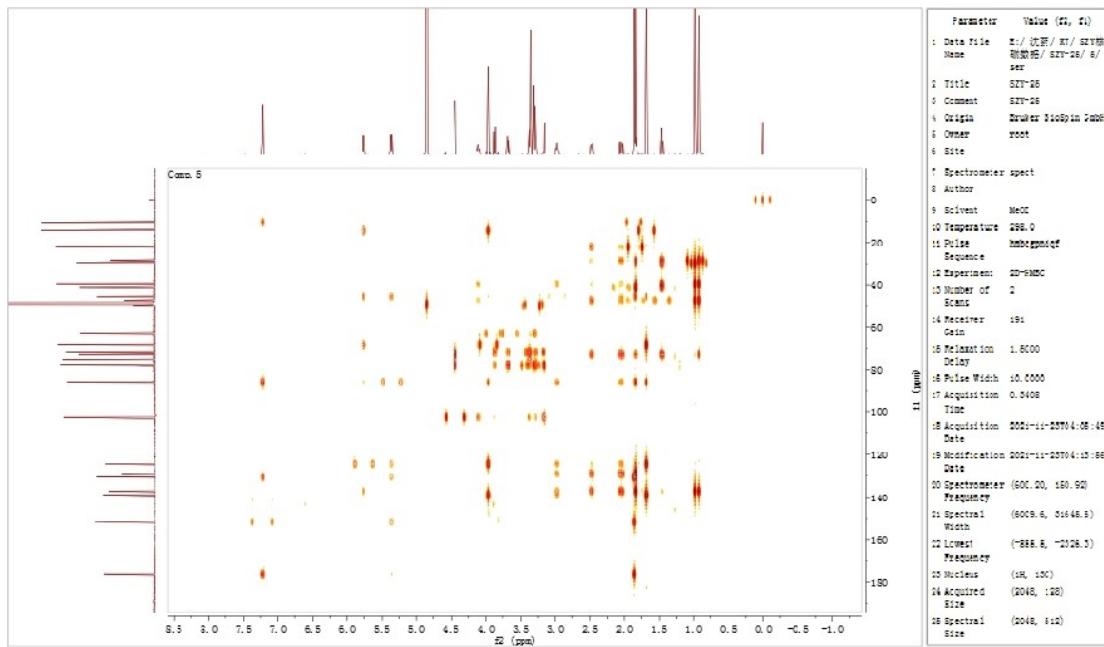


Fig S 30 The HMBC spectrum (600 MHz, Methanol-*d*₄) of compound 5.

SUPPORTING INFORMATION

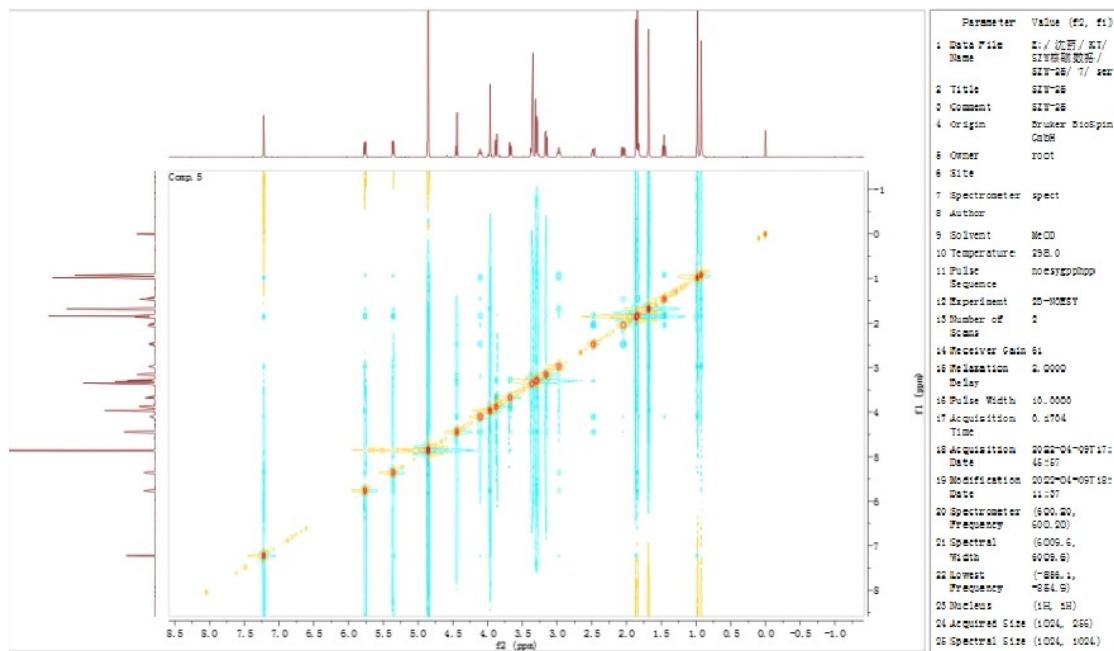


Fig S 31 The NOESY spectrum (600 MHz, $\text{MeOH}-d_4$) of compound 5.

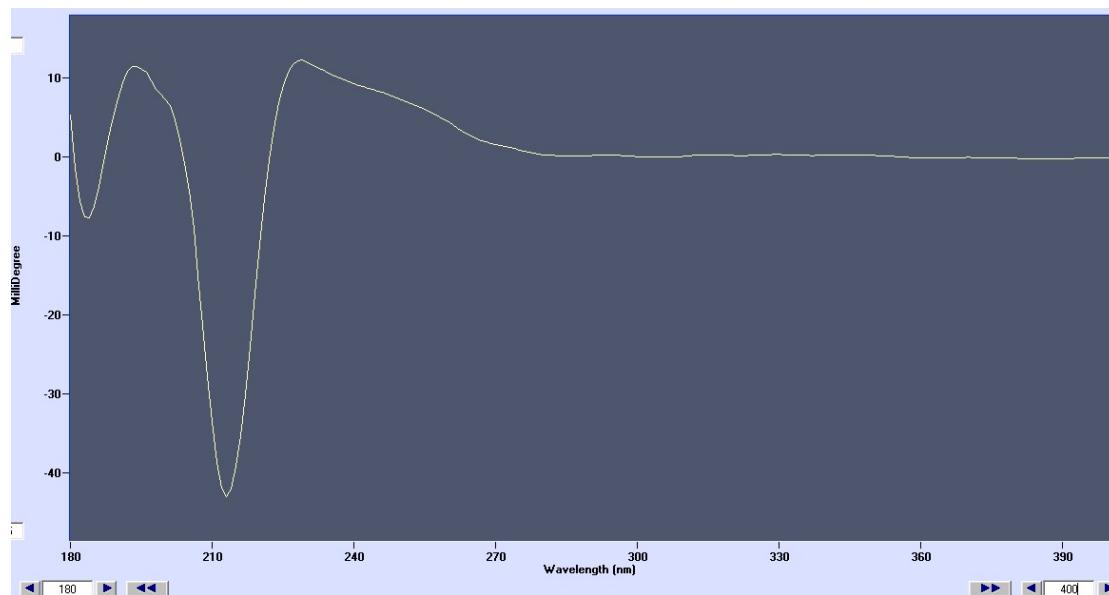
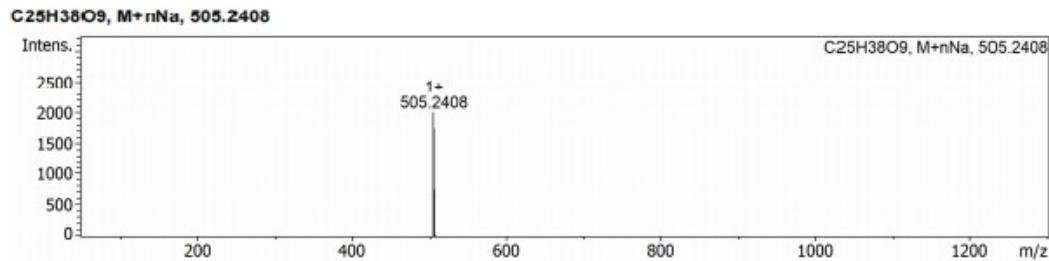
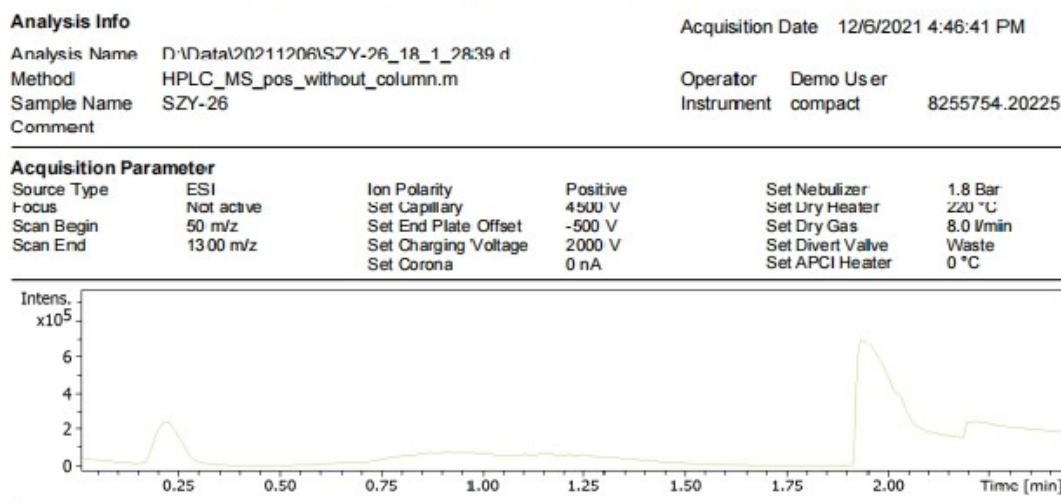


Fig S 32 The CD spectrum (MeOH) of compound 5'.

SUPPORTING INFORMATION

Compound Spectrum SmartFormula Report



SUPPORTING INFORMATION

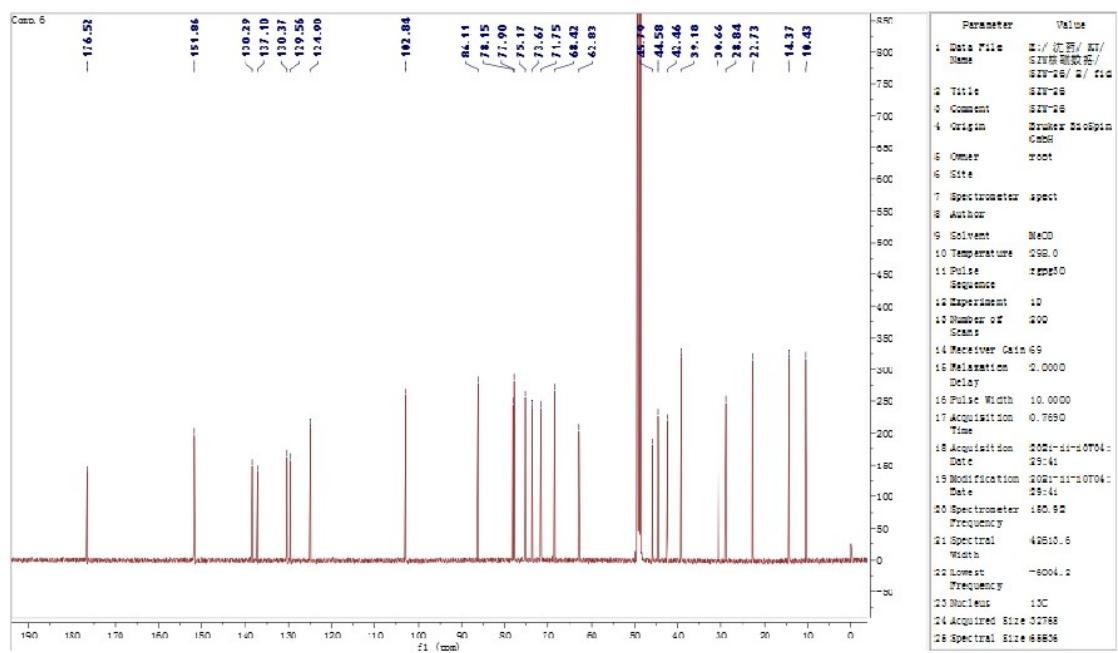


Fig S 35 The ^{13}C NMR spectrum (150 MHz, Methanol- d_4) of compound 6.

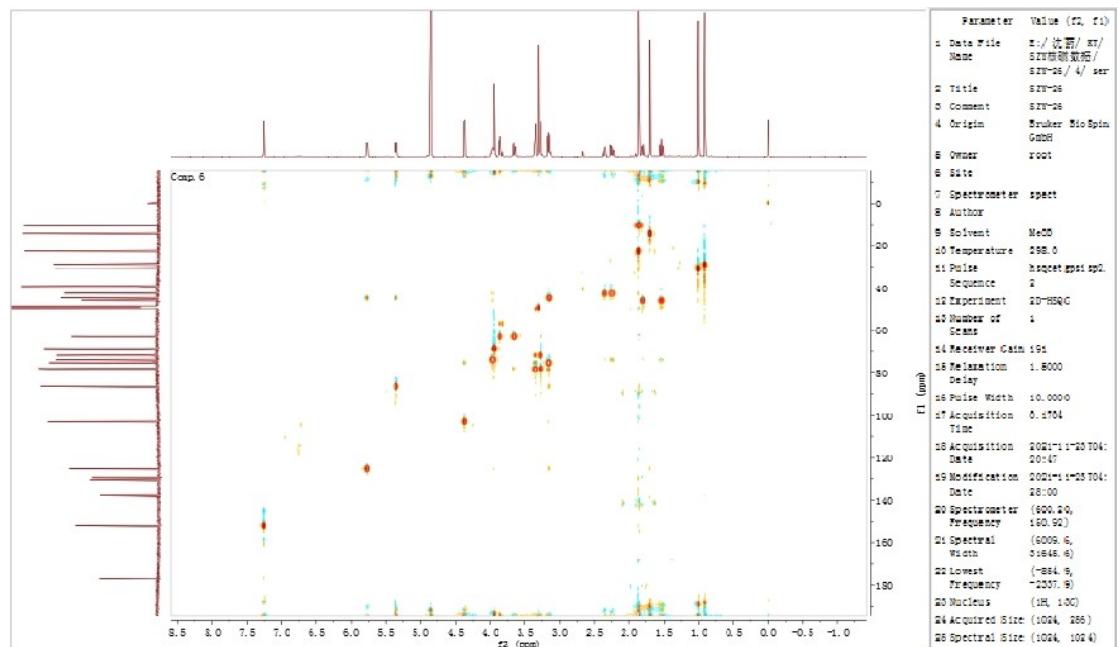


Fig S 36 The HSQC spectrum (600 MHz, Methanol- d_4) of compound 6.

SUPPORTING INFORMATION

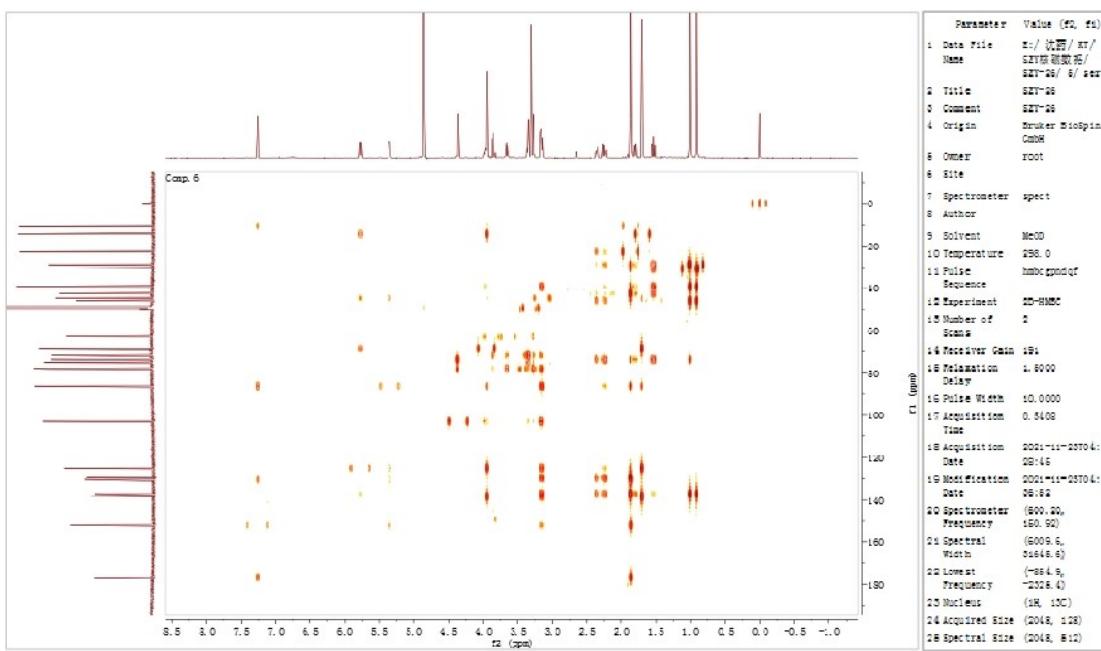


Fig S 37 The HMBC spectrum (600 MHz, Methanol- d_4) of compound 6.

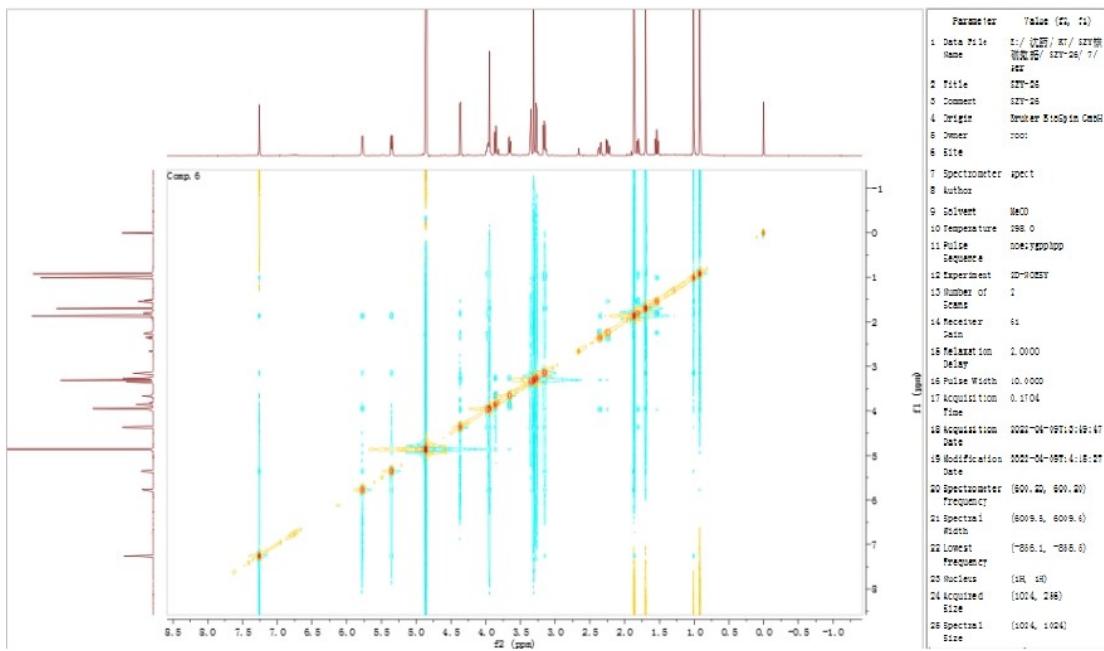


Fig S 38 The NOESY spectrum (600 MHz, Methanol- d_4) of compound 6.

SUPPORTING INFORMATION

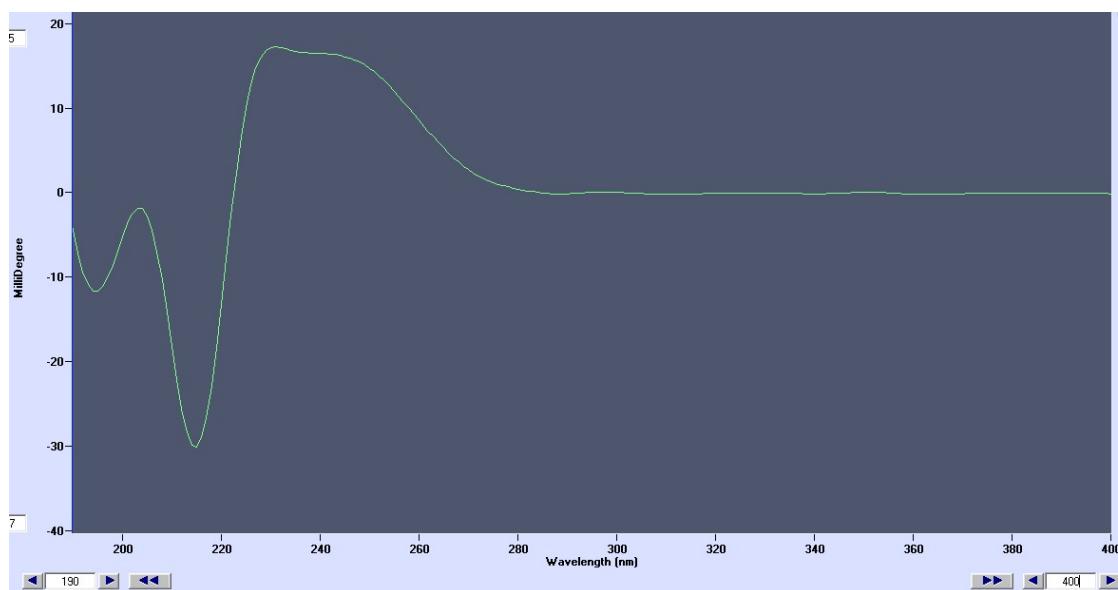


Fig S 39 The CD spectrum (MeOH) of compound 6'.

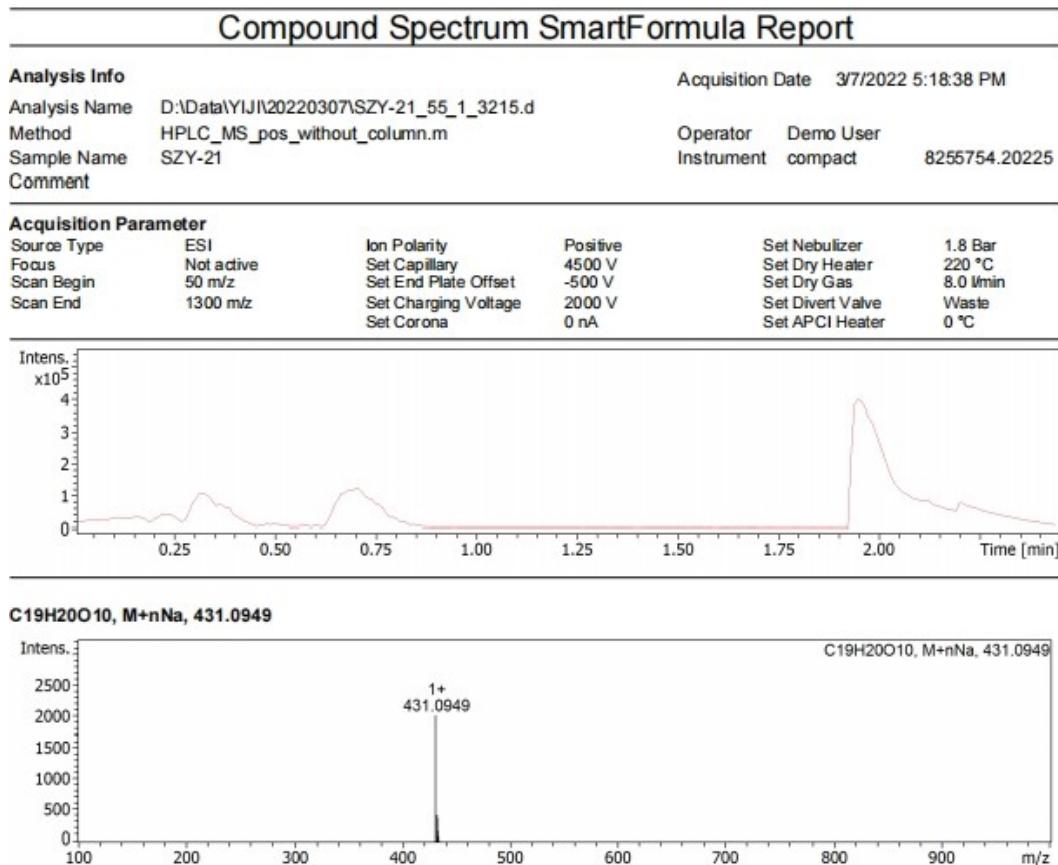


Fig S 40 The HRESIMS spectrum of compound 7.

SUPPORTING INFORMATION

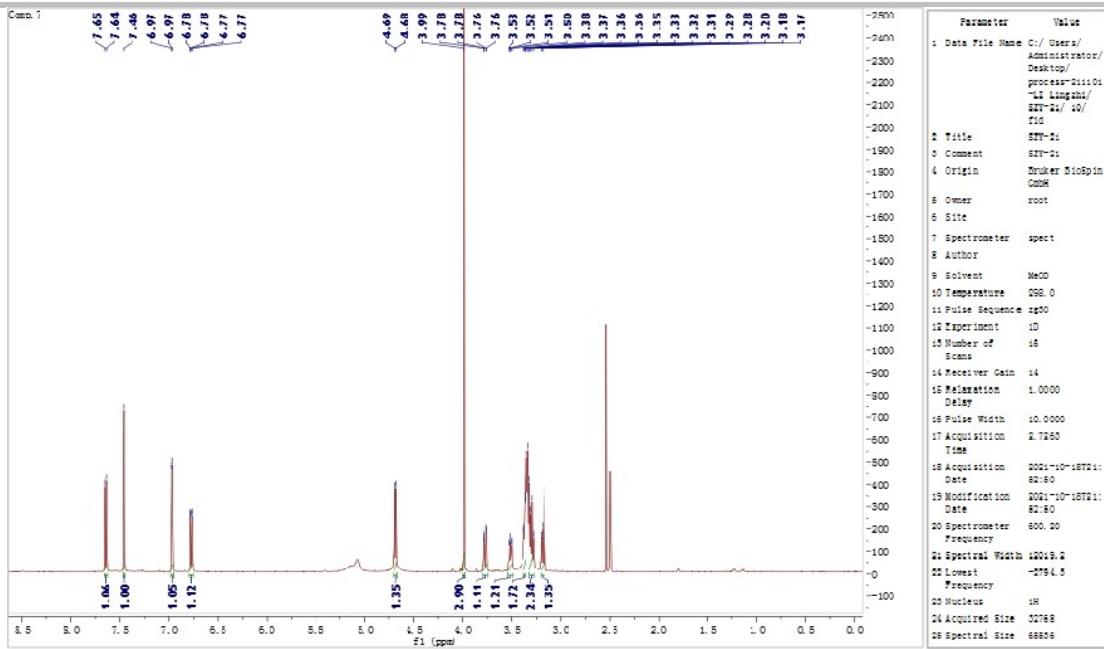


Fig S 41 The ^1H NMR spectrum (600 MHz, DMSO- d_6) of compound 7.

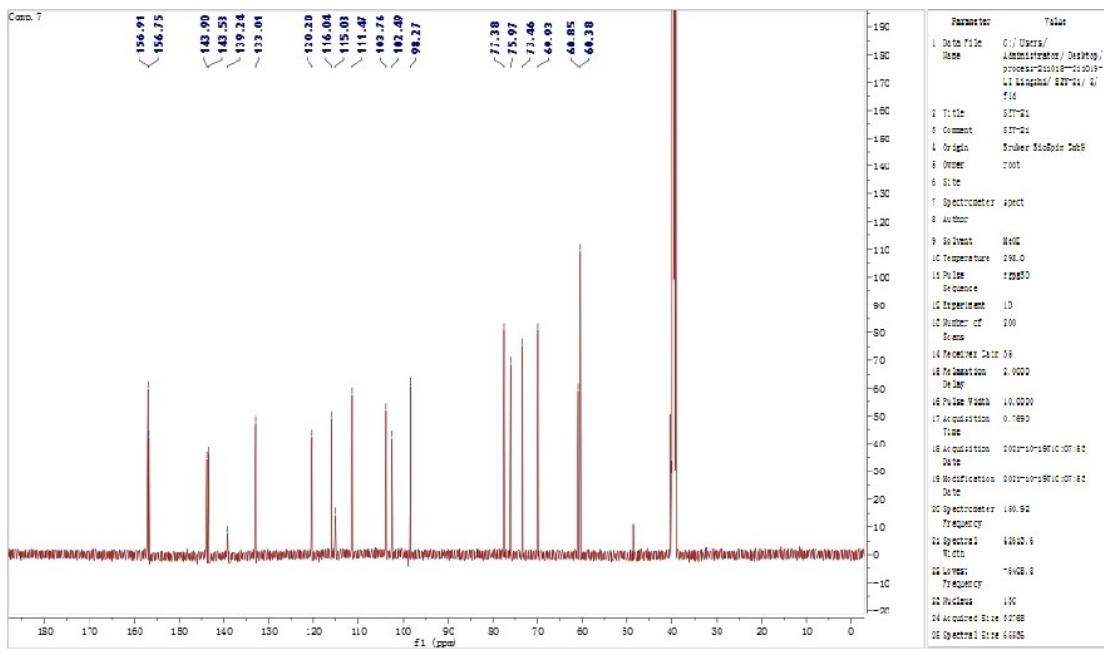


Fig S 42 The ^{13}C NMR spectrum (150 MHz, DMSO- d_6) of compound 7.

SUPPORTING INFORMATION

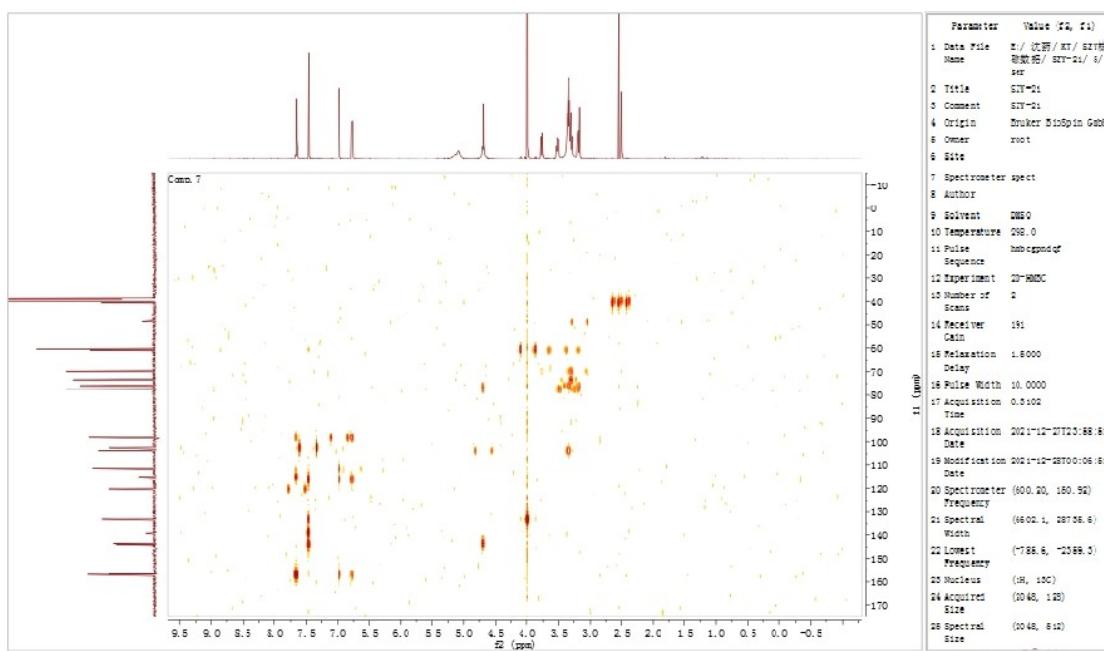


Fig S 43 The HMBC spectrum (600 MHz, DMSO-*d*₆) of compound 7.

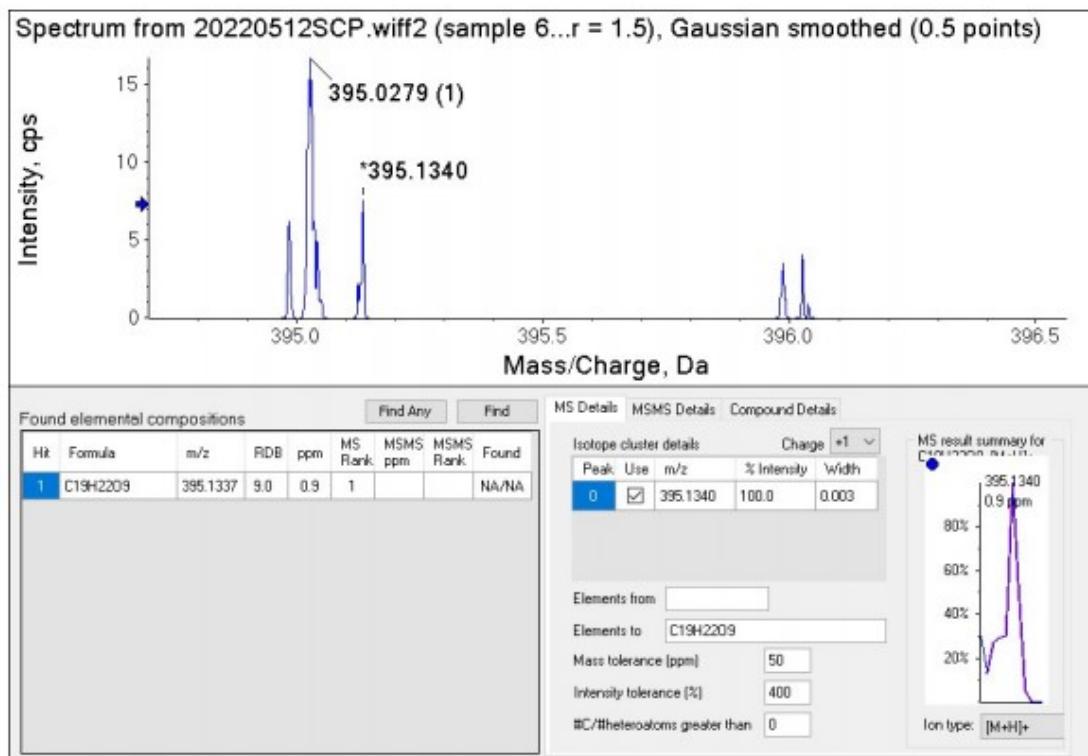


Fig S 44 The HRESIMS spectrum of compound 8.

SUPPORTING INFORMATION

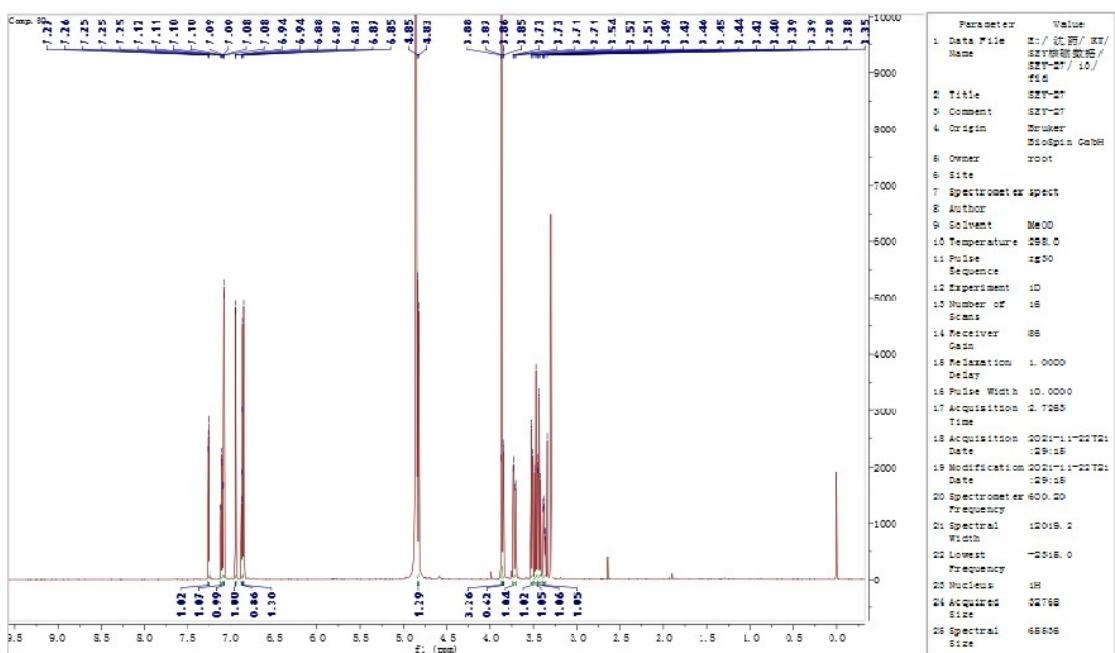


Fig S 45 The ¹H NMR spectrum (600 MHz, Methanol-*d*₄) of compound 8.

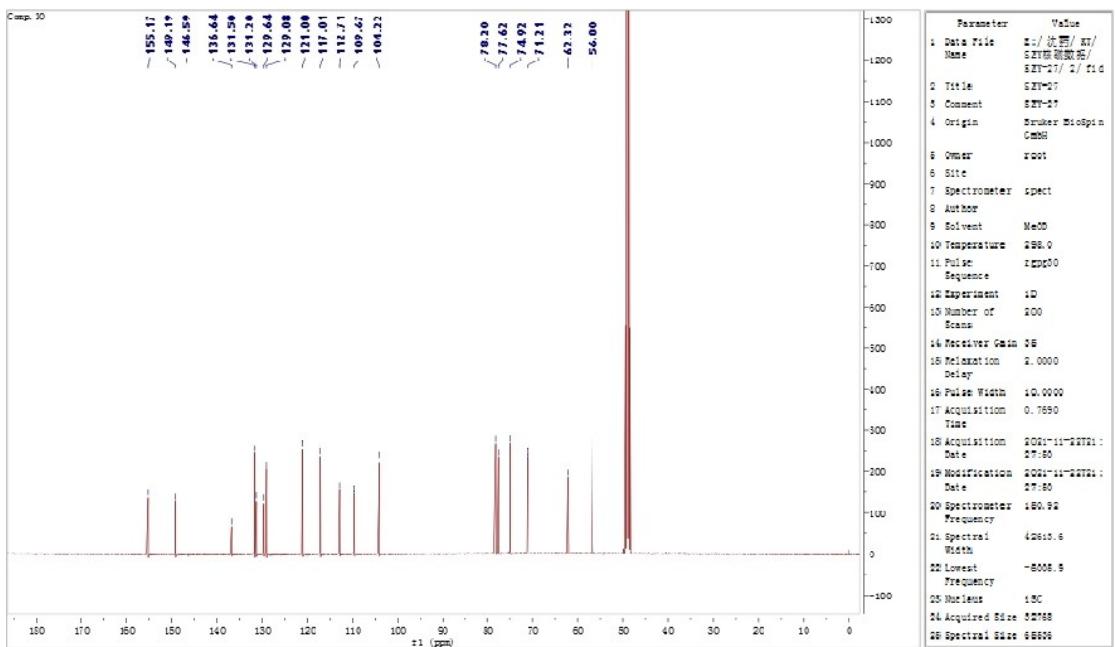


Fig S 46 The ¹³C NMR spectrum (150 MHz, Methanol-*d*₄) of compound 8.

SUPPORTING INFORMATION

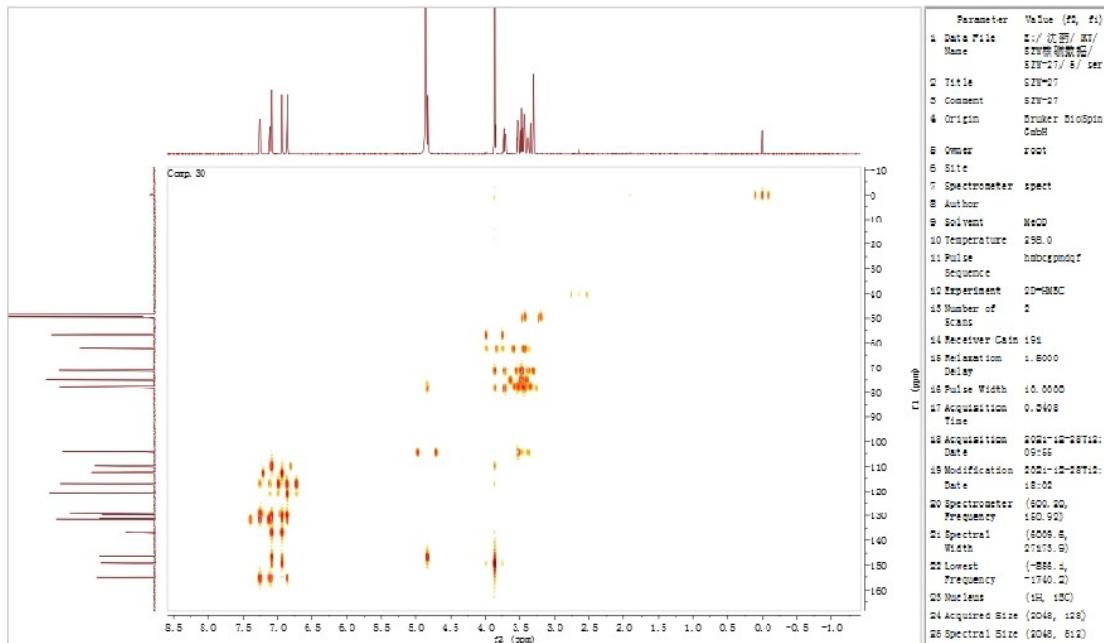


Fig S 47 The HMBC spectrum (600 MHz, Methanol-*d*₄) of compound 8.

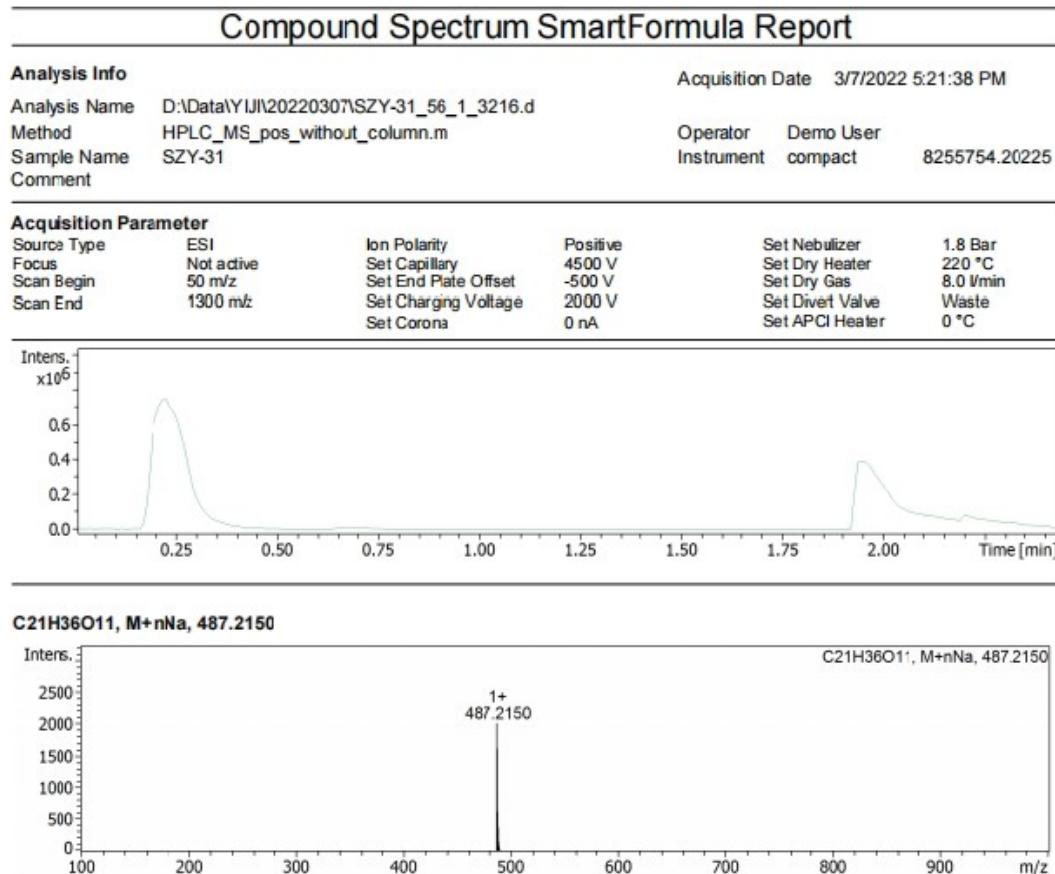


Fig S 48 The HRESIMS spectrum of compound 9.

SUPPORTING INFORMATION

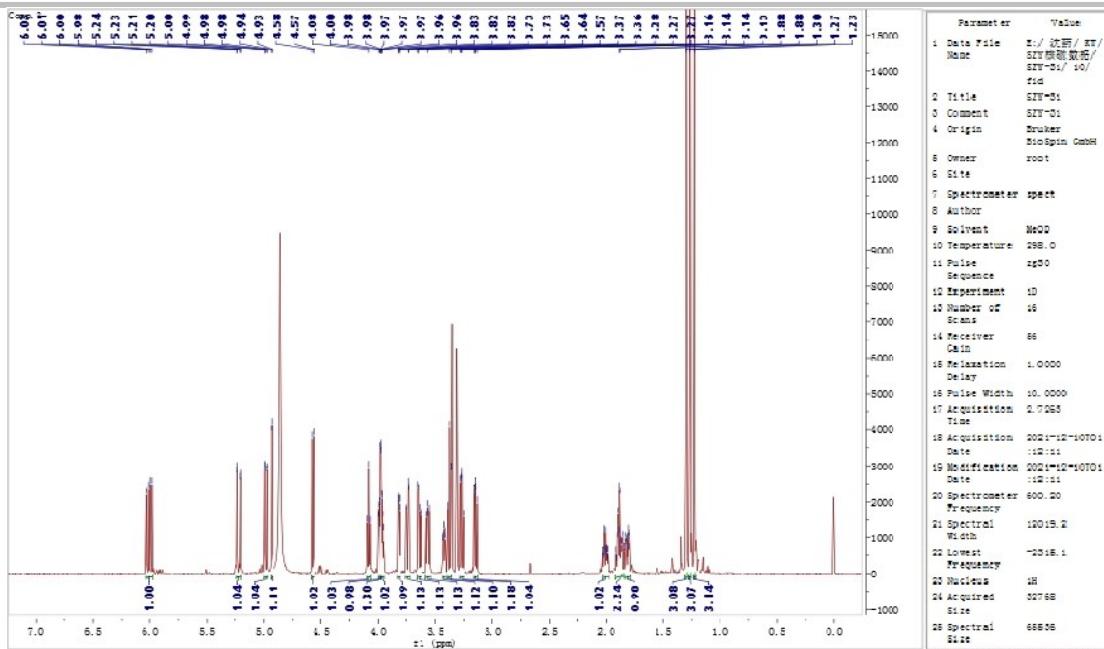


Fig S 49 The ¹H NMR spectrum (600 MHz, Methanol-*d*₄) of compound 9.

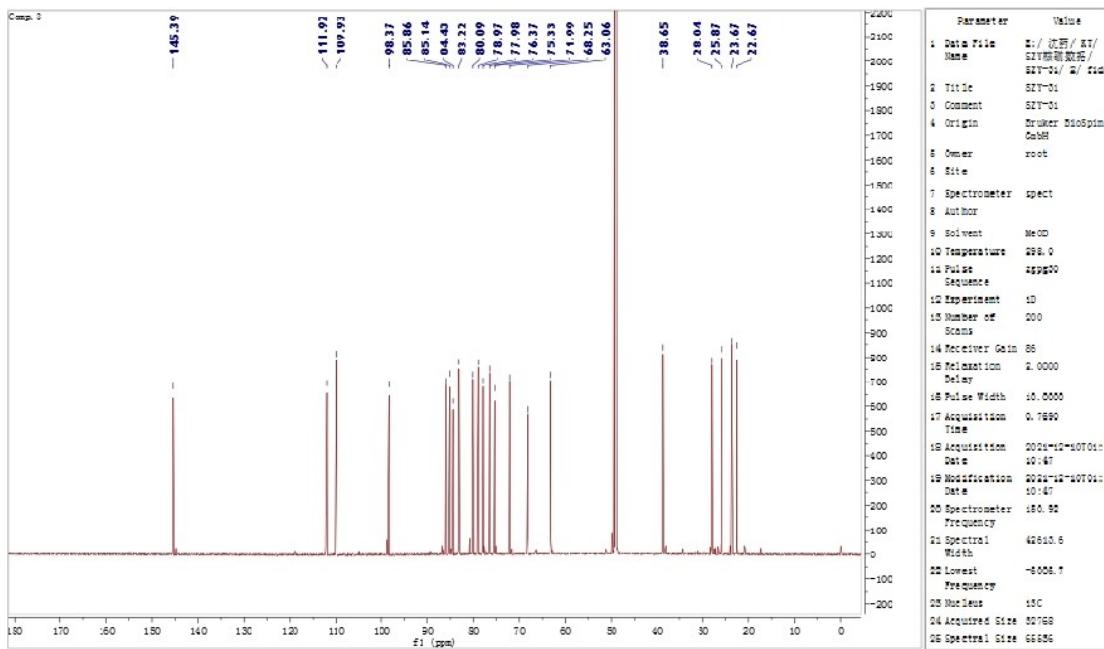


Fig S 50 The ¹³C NMR spectrum (150 MHz, Methanol-*d*₄) of compound 9.

SUPPORTING INFORMATION

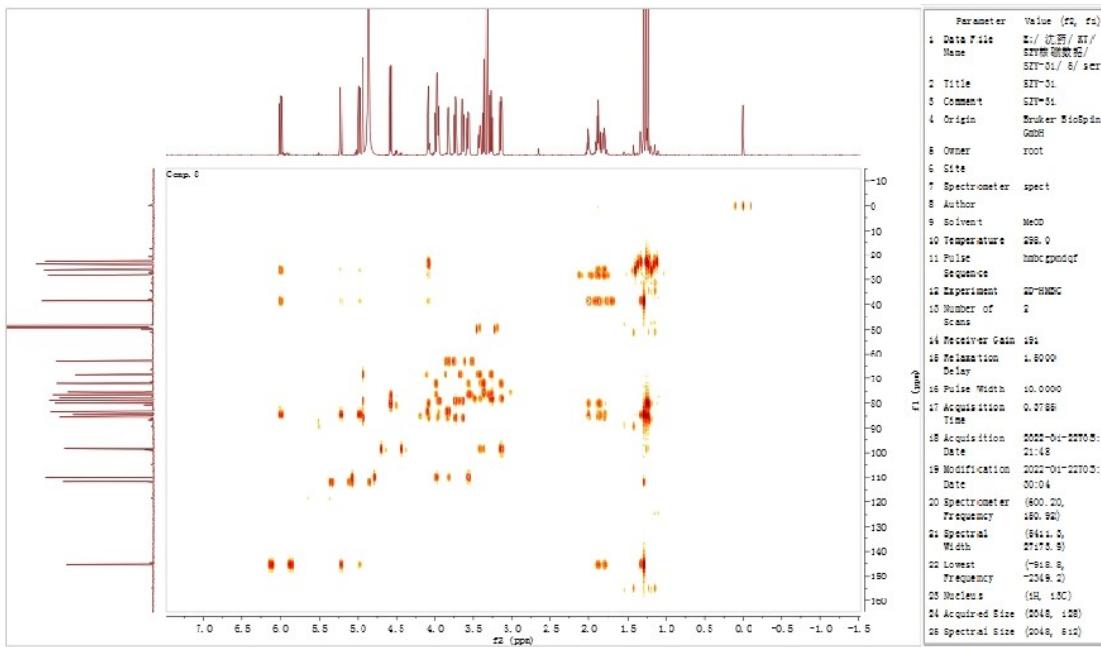


Fig S 51 The HMBC spectrum (600 MHz, Methanol-*d*₄) of compound 9.

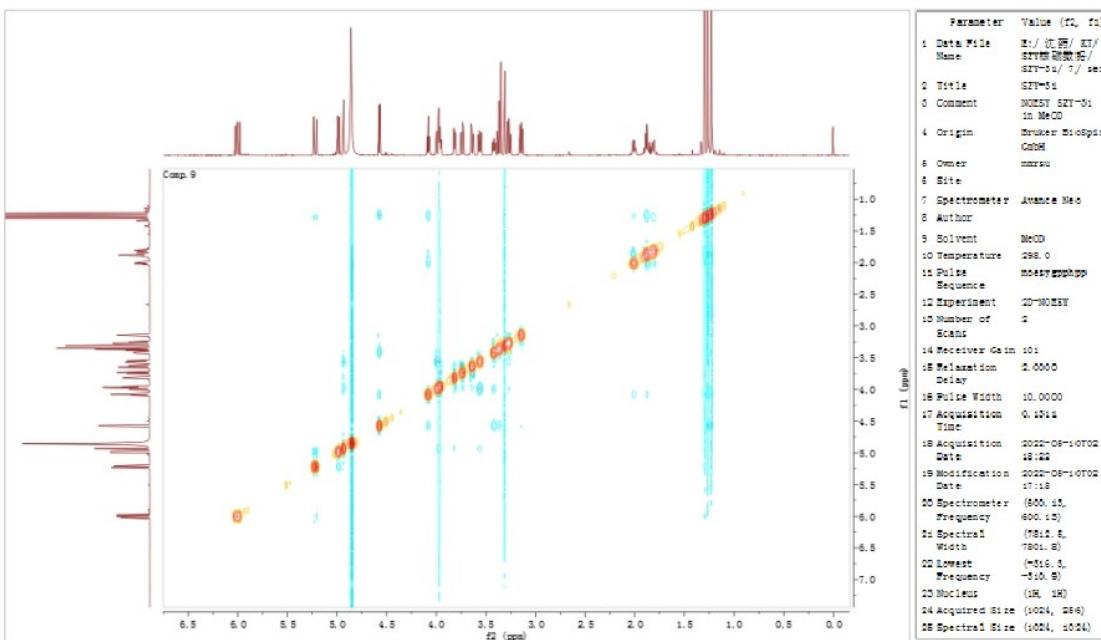


Fig S 52 The NOESY spectrum (600 MHz, Methanol-*d*₄) of compound 9.

SUPPORTING INFORMATION

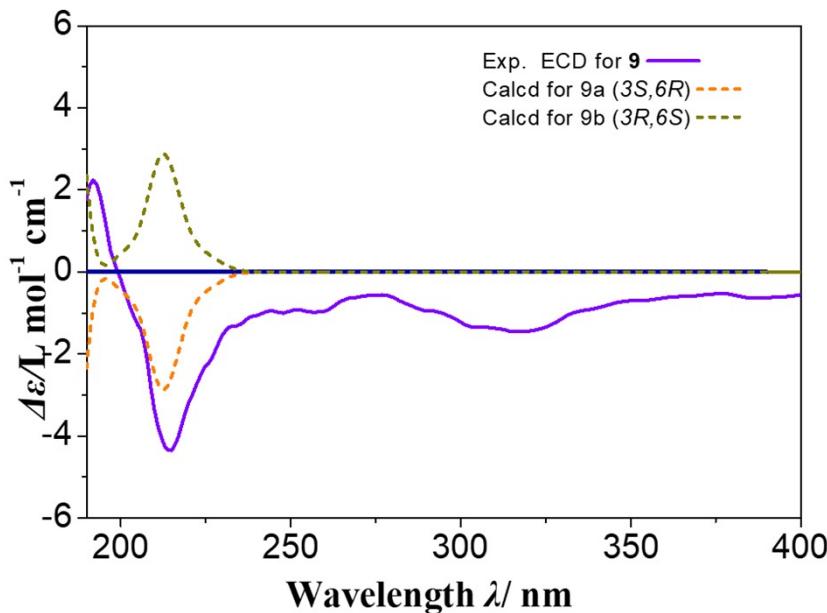


Fig S 53 The CD spectrum (MeOH) of compound **9**.

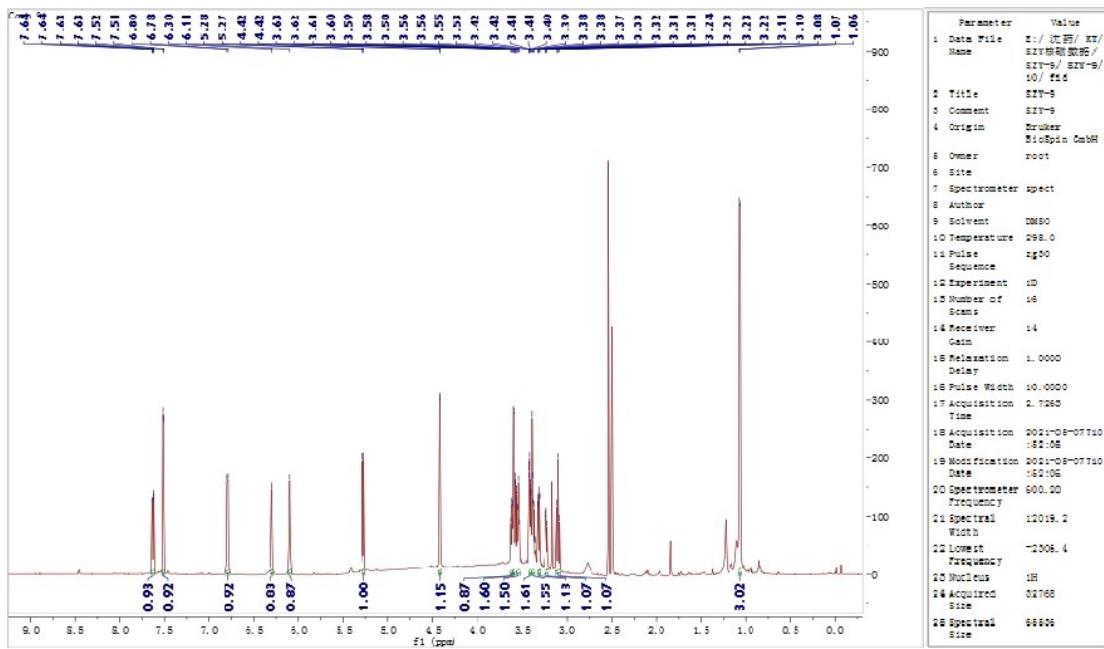


Fig S 54 The ^1H NMR spectrum (600 MHz, $\text{DMSO}-d_6$) of compound **10**.

SUPPORTING INFORMATION

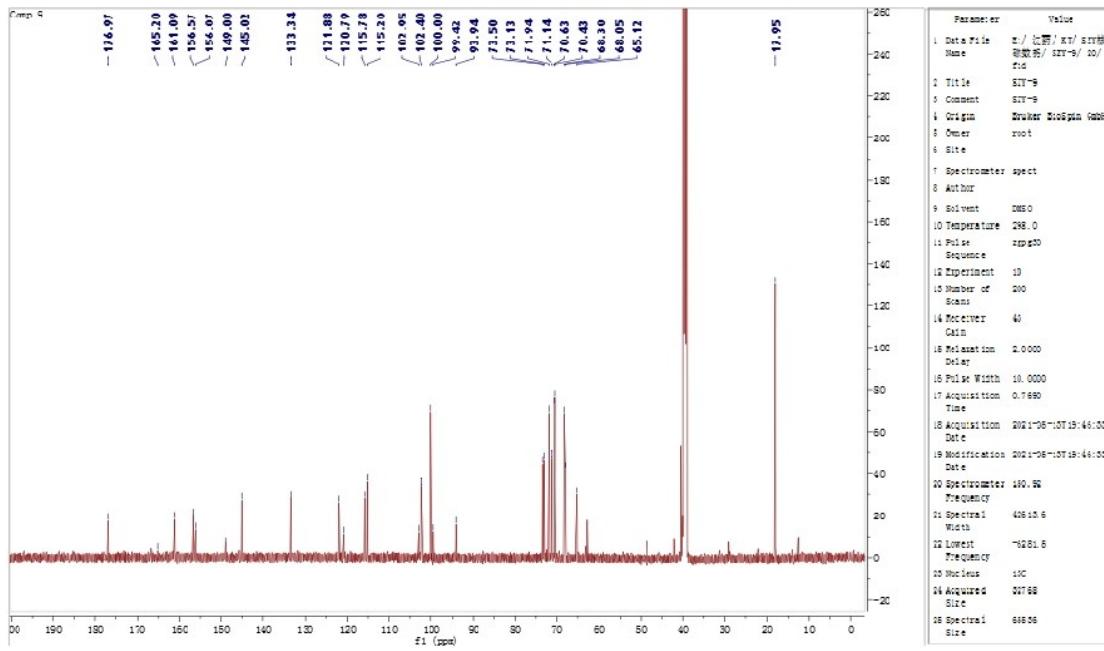


Fig S 55 The ^{13}C NMR spectrum (150 MHz, $\text{DMSO}-d_6$) of compound **10**.

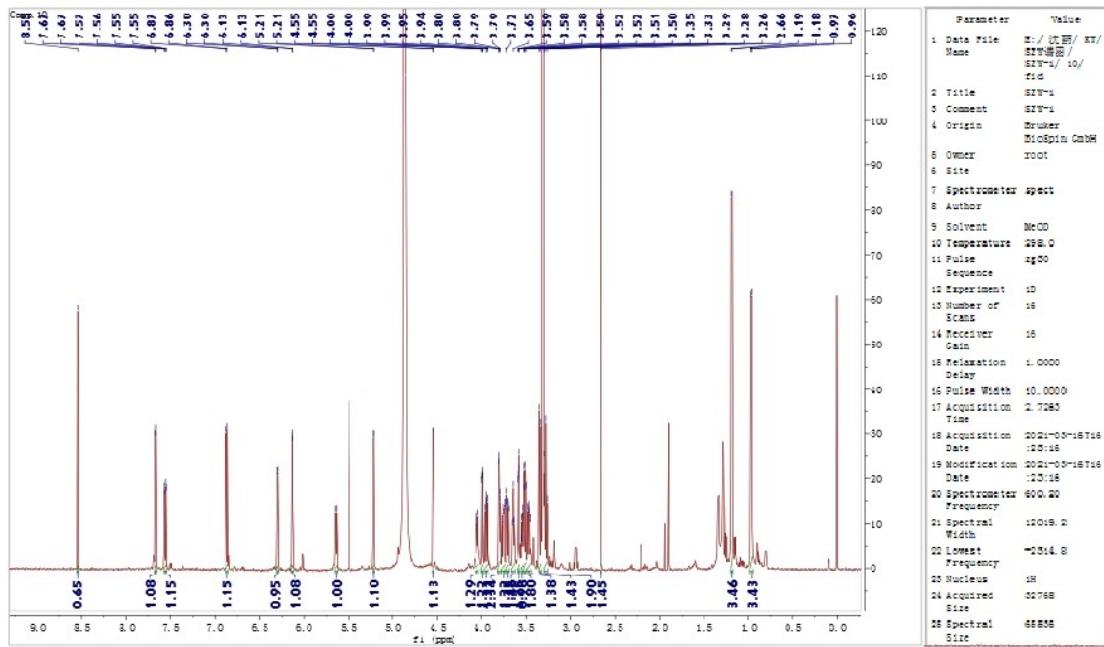


Fig S 56 The ^1H NMR spectrum (600 MHz, Methanol- d_4) of compound 11.

SUPPORTING INFORMATION

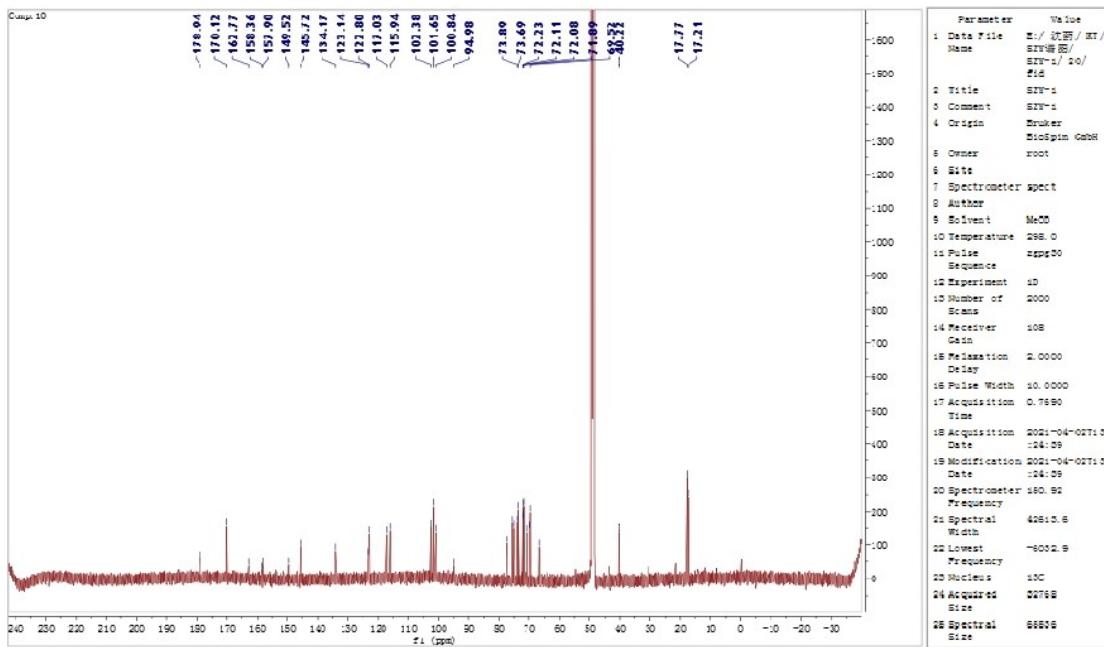


Fig S 57 The ¹³C NMR spectrum (150 MHz, Methanol-*d*₄) of compound 11.

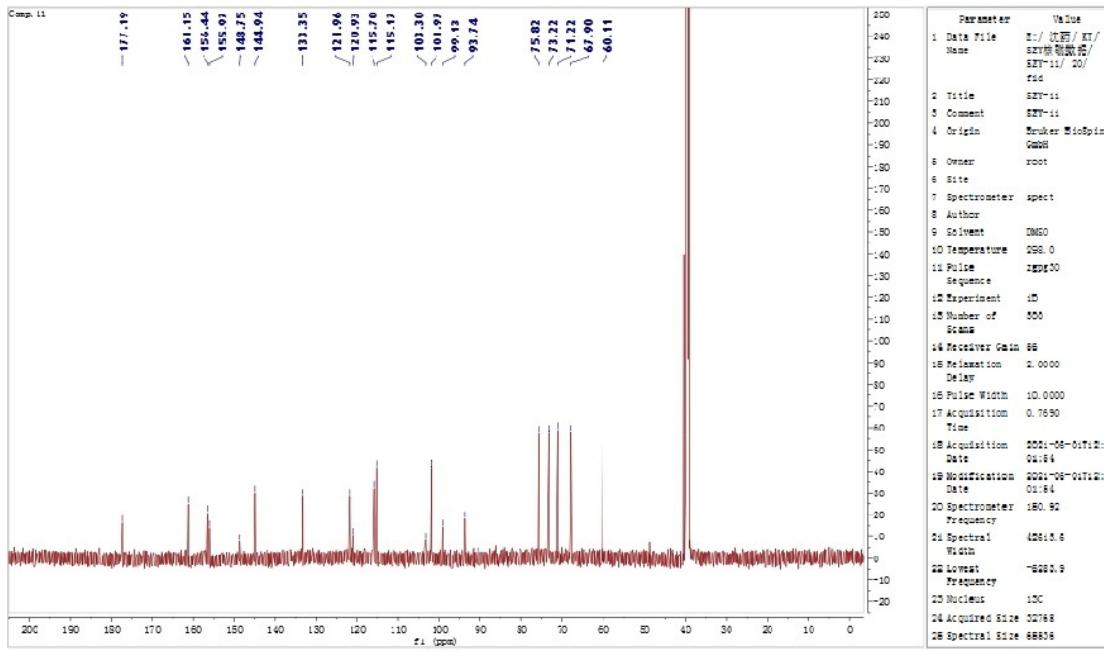


Fig S 58 The ¹³C NMR spectrum (150 MHz, DMSO-*d*₆) of compound 12.

SUPPORTING INFORMATION

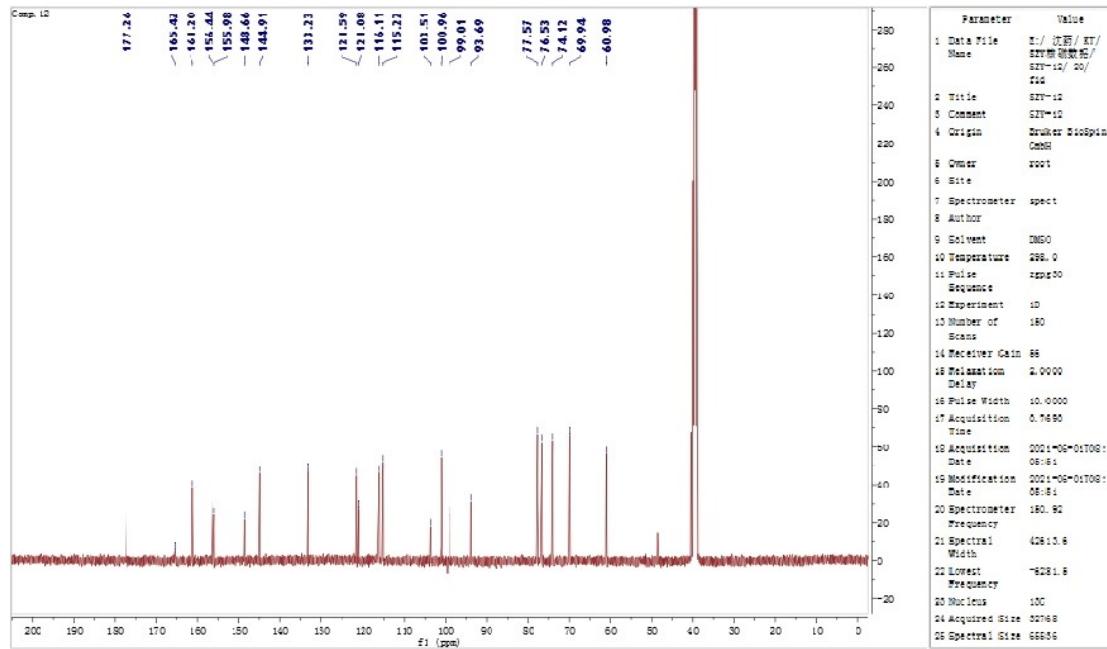


Fig S 59 The ^{13}C NMR spectrum (150 MHz, $\text{DMSO}-d_6$) of compound 13.

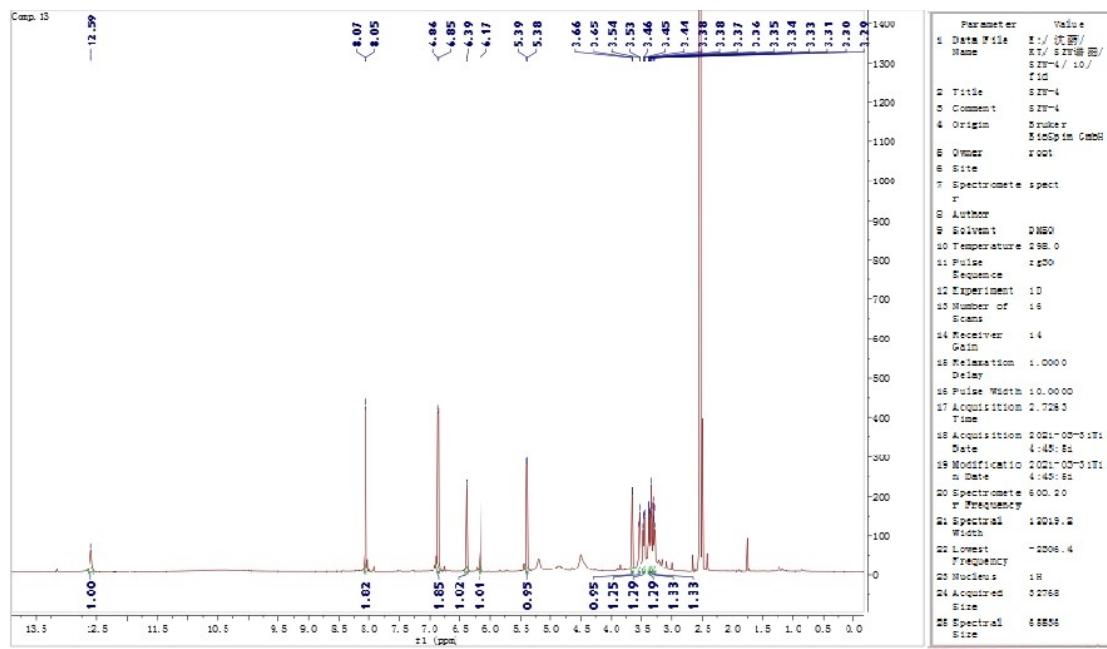


Fig S 60 The ^1H NMR spectrum (600 MHz, $\text{DMSO}-d_6$) of compound 14.

SUPPORTING INFORMATION

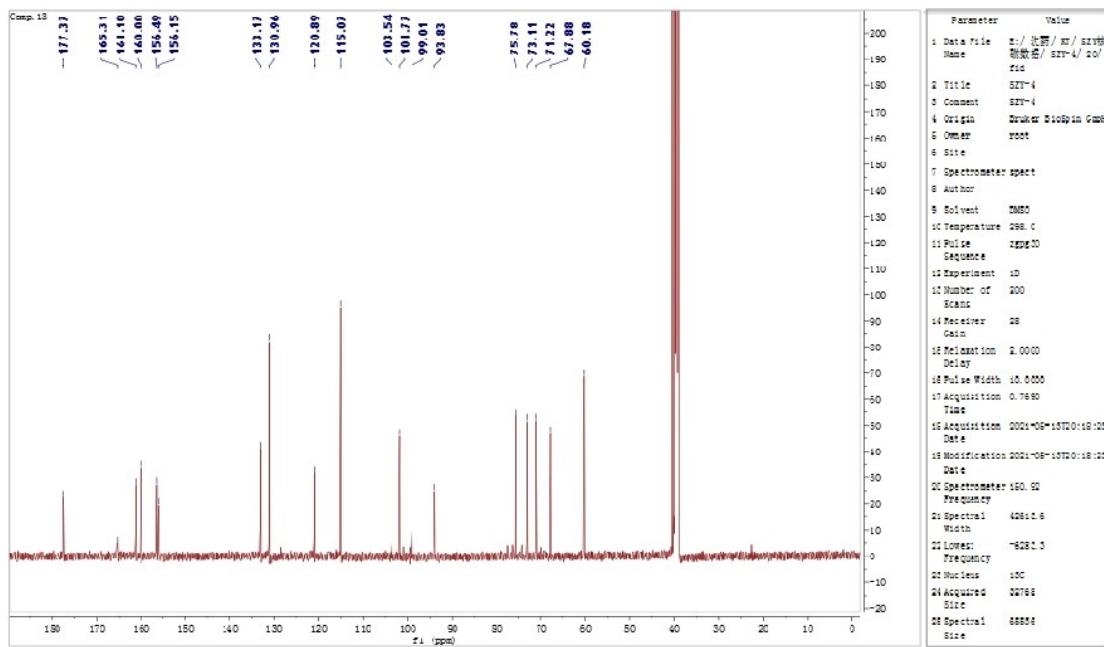


Fig S 61 The ¹³C NMR spectrum (150 MHz, DMSO-*d*₆) of compound 14.

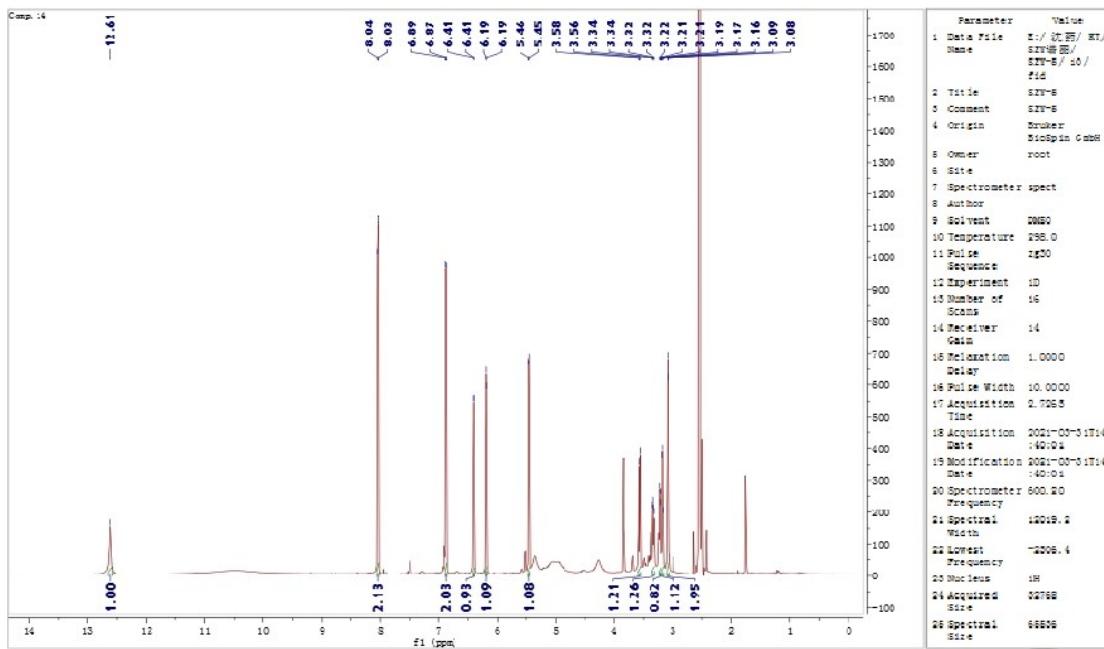


Fig S 62 The ¹H NMR spectrum (600 MHz, DMSO-*d*₆) of compound 15.

SUPPORTING INFORMATION

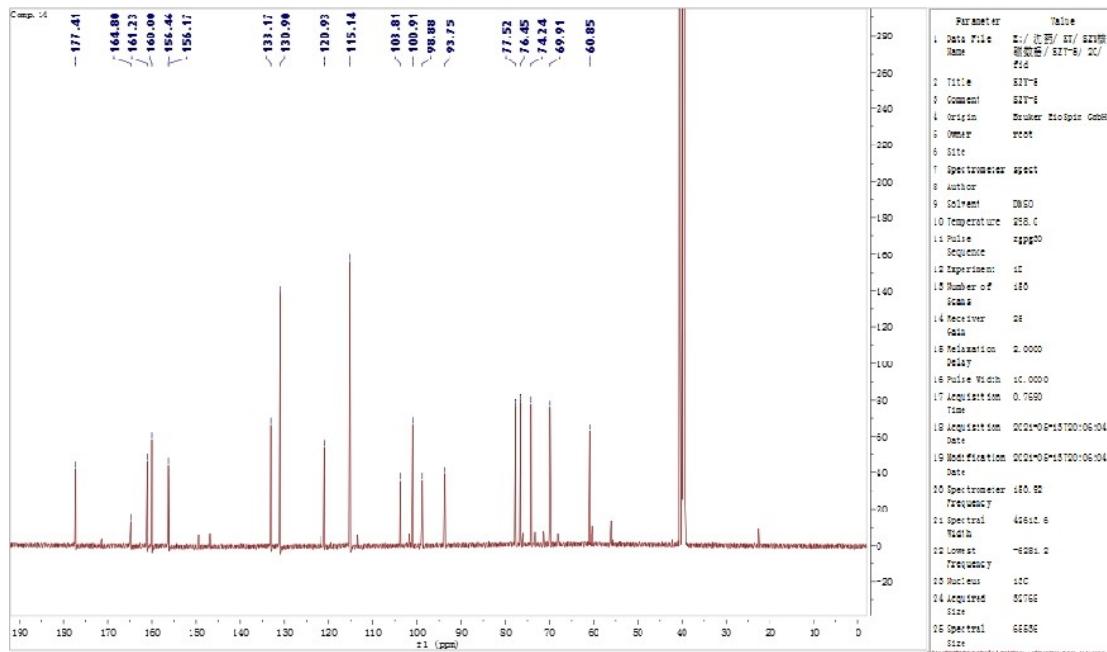


Fig S 63 The ¹³C NMR spectrum (150 MHz, DMSO-*d*₆) of compound 15.

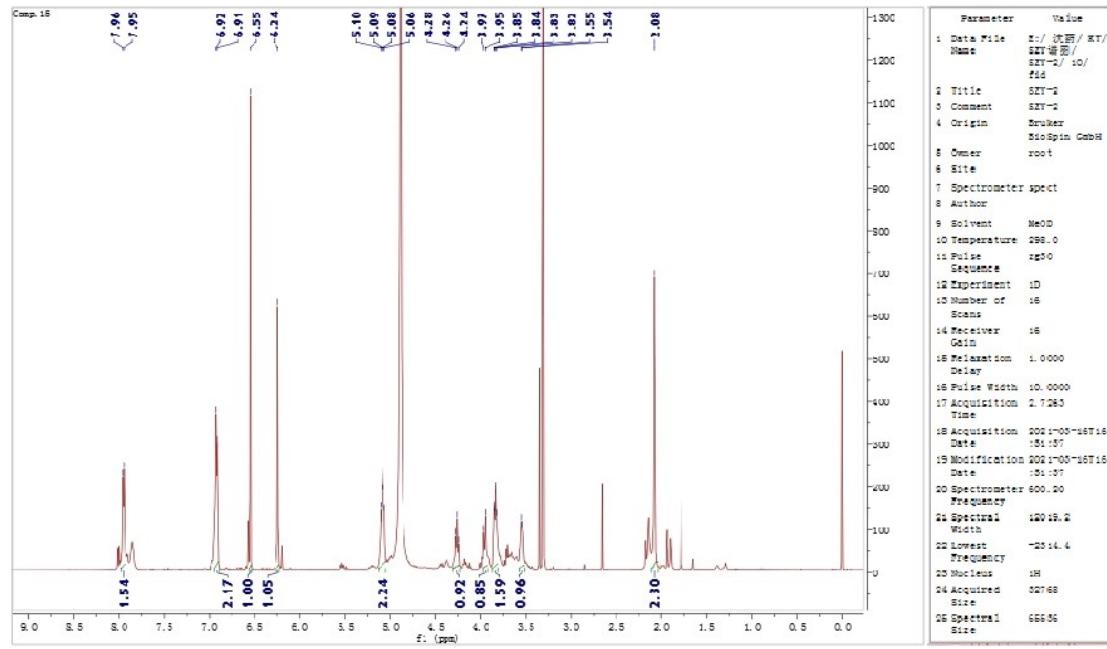


Fig S 64 The ¹H NMR spectrum (600 MHz, DMSO-*d*₆) of compound 16.

SUPPORTING INFORMATION

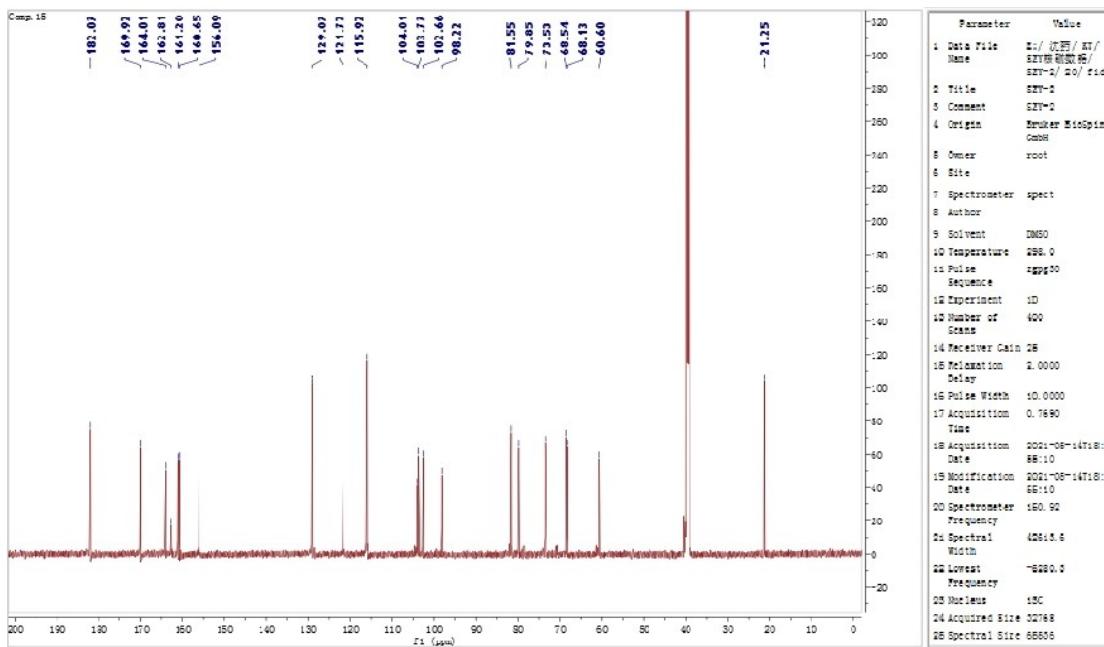


Fig S 65 The ¹³C NMR spectrum (150 MHz, DMSO-*d*₆) of compound 16.

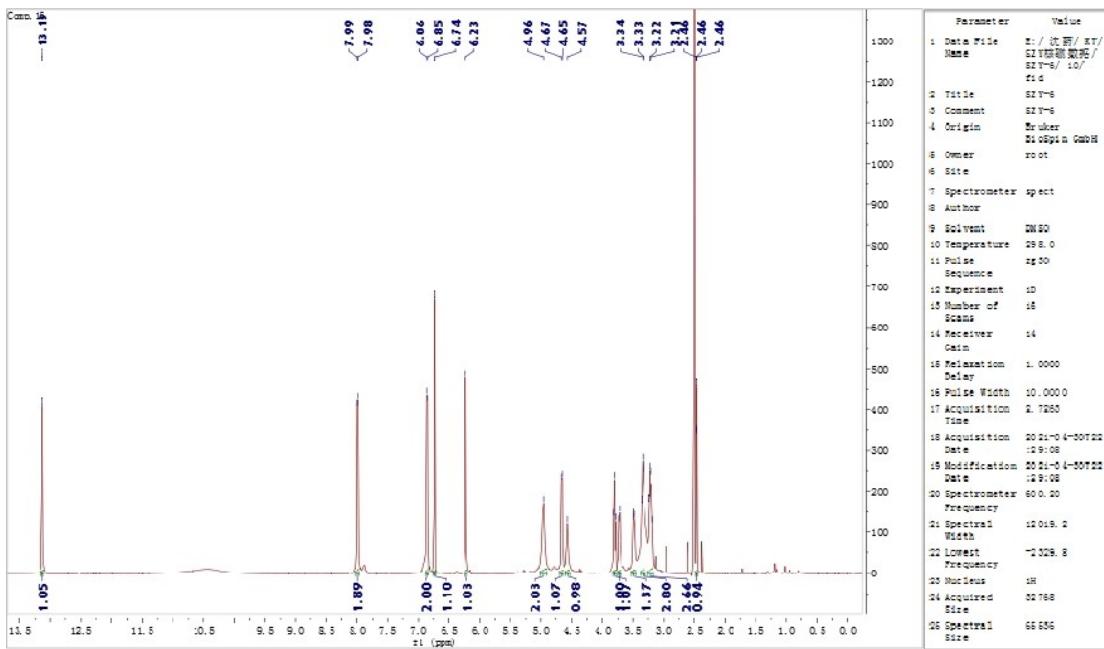


Fig S 66 The ¹H NMR spectrum (600 MHz, DMSO-*d*₆) of compound 17.

SUPPORTING INFORMATION

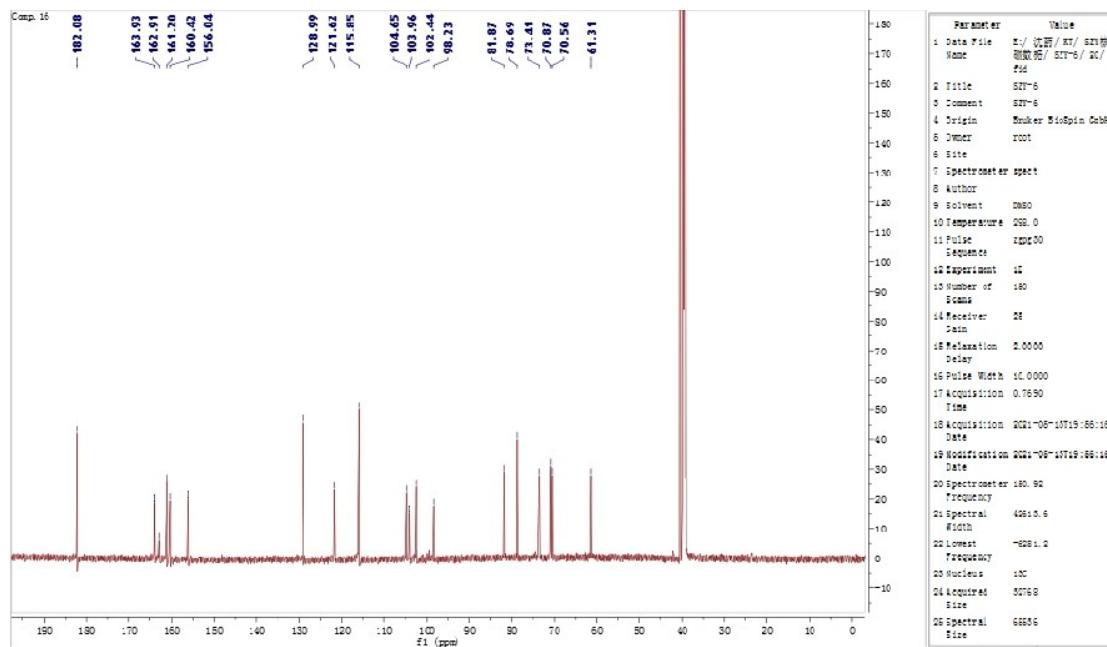


Fig S 67 The ¹³C NMR spectrum (150 MHz, DMSO-*d*₆) of compound 17.

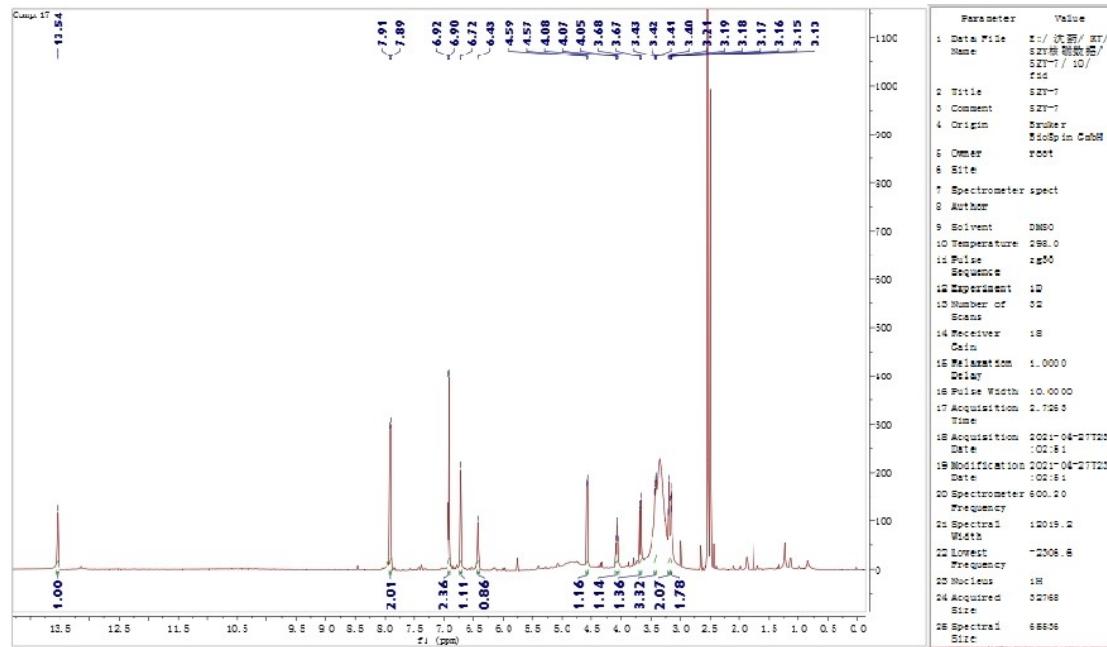


Fig S 68 The ¹H NMR spectrum (600 MHz, DMSO-*d*₆) of compound 18.

SUPPORTING INFORMATION

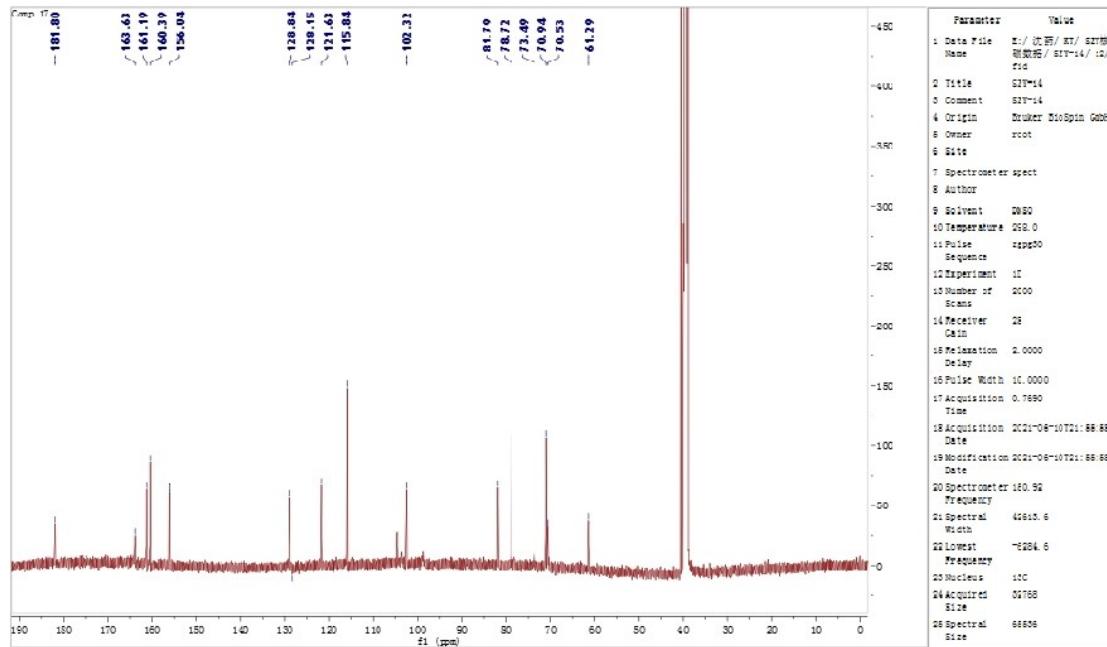


Fig S 69 The ^{13}C NMR spectrum (150 MHz, DMSO- d_6) of compound **18**.

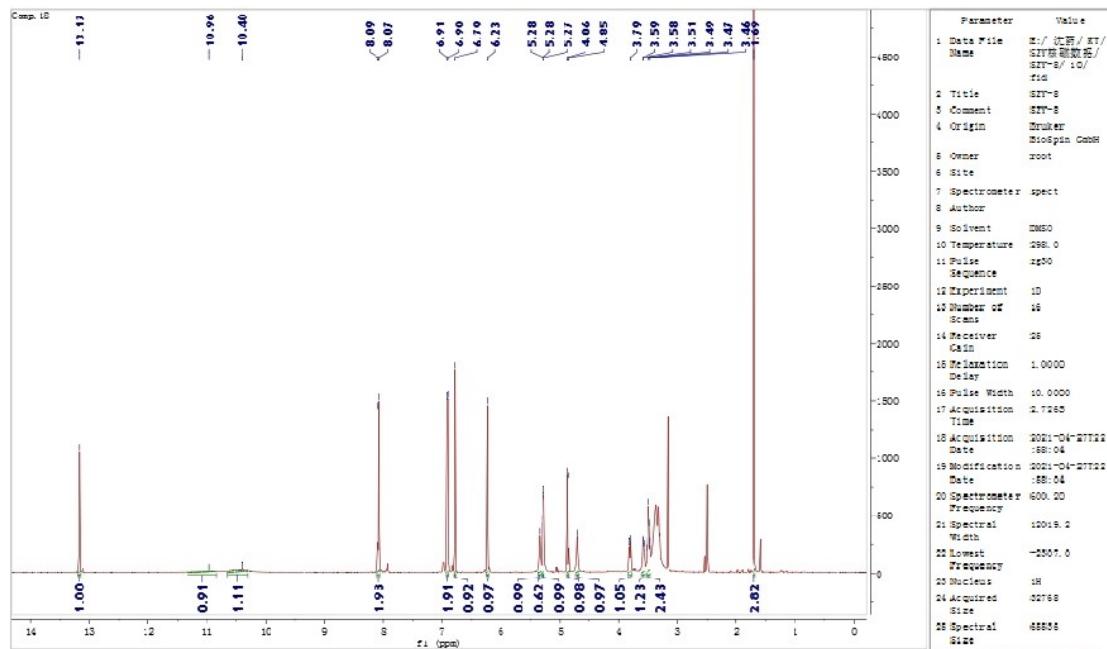


Fig S 70 The ^1H NMR spectrum (600 MHz, DMSO- d_6) of compound **19**.

SUPPORTING INFORMATION

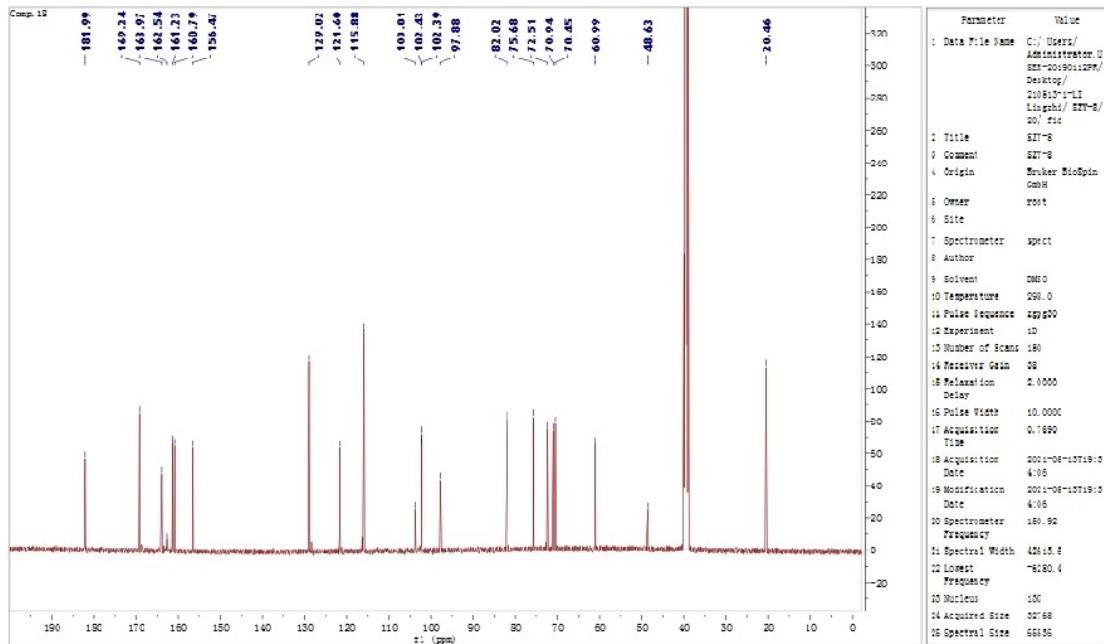


Fig S 71 The ^{13}C NMR spectrum (150 MHz, DMSO- d_6) of compound 19.

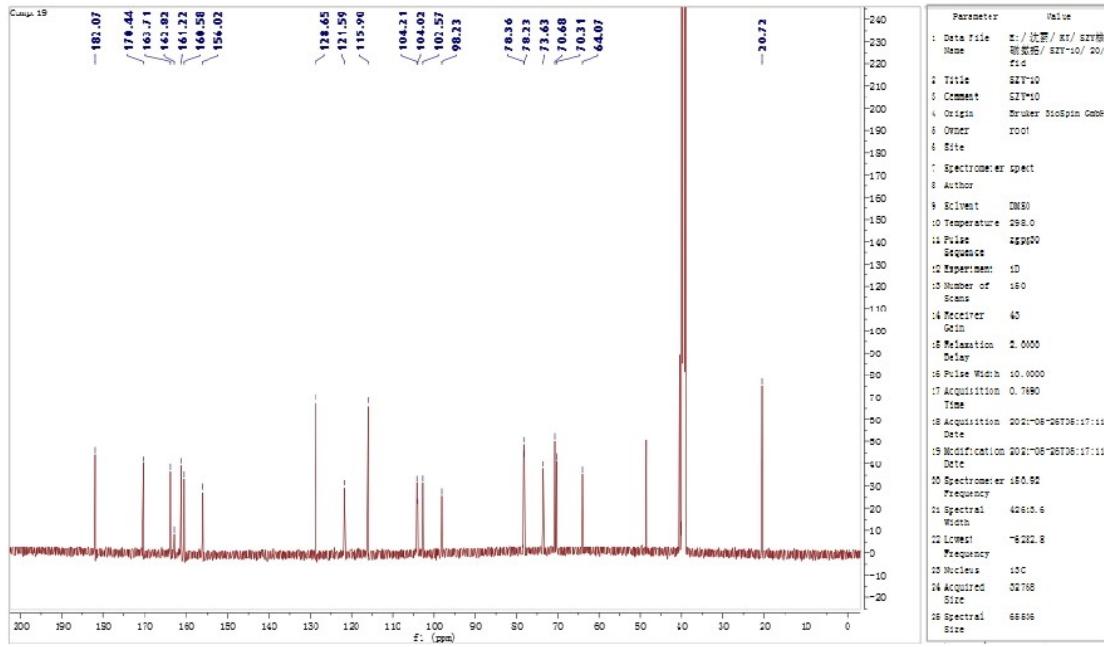


Fig S 72 The ^{13}C NMR spectrum (150 MHz, DMSO- d_6) of compound 20.

SUPPORTING INFORMATION

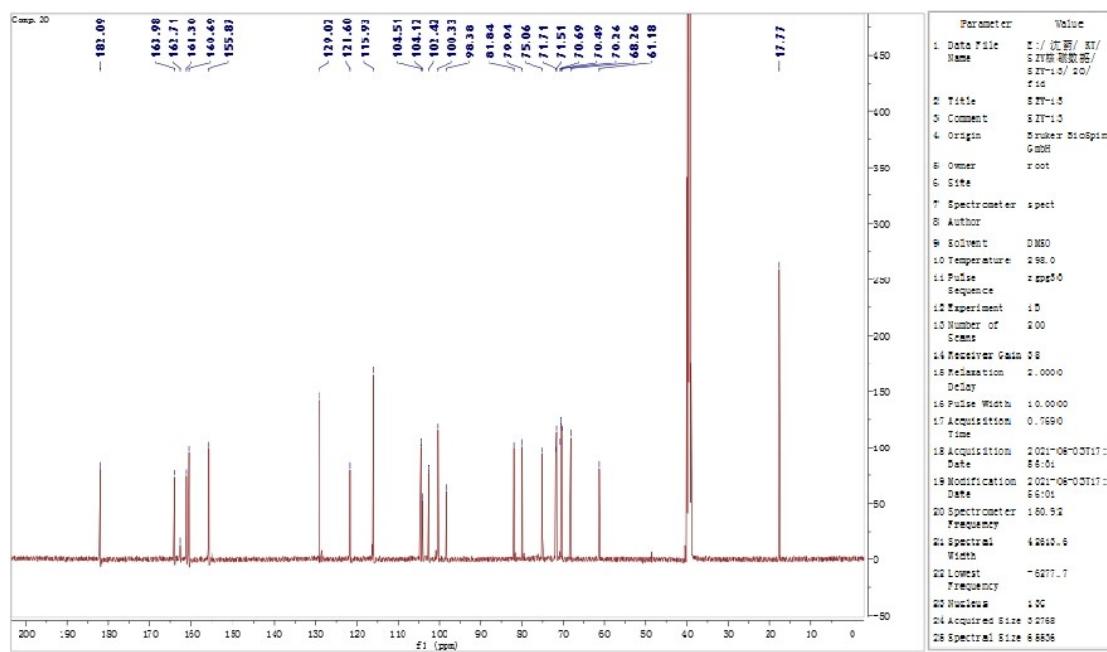


Fig S 73 The ^{13}C NMR spectrum (150 MHz, DMSO- d_6) of compound 21.

SUPPORTING INFORMATION

Table S 1 ^1H (600 MHz) and ^{13}C (150 MHz) data of compounds **1** and **2** (Methanol- d_4).

No.	1		2	
	δ_{H} , mult, J (Hz)	δ_{C}	δ_{H} , mult, J (Hz)	δ_{C}
1		173.1		173.1
2	2.76 (1H, dd, $J = 18.0, 1.2$ Hz) 2.71 (1H, dd, $J = 18.0, 4.8$ Hz)	36.7	2.74 (1H, dd, $J = 18.0, 1.8$ Hz) 2.69 (1H, dd, $J = 18.0, 4.8$ Hz)	36.6
3	4.26 (1H, m) 2.29 (1H, 1H, m)	73.2	4.18 (1H, m) 2.18 (1H, m)	73.1
4	1.68 (1H, ddd, $J = 14.1, 11.4, 2.6$ Hz)	36.8	1.60 (1H, ddd, $J = 14.2, 11.4, 2.8$ Hz)	36.7
5	4.81 (1H, m)	74.8	4.78 (1H, m)	74.8
6	1.31 (3H, d, $J = 6.4$ Hz) Glc	21.6	1.25 (3H, d, $J = 6.4$ Hz) Glc	21.6 Glc
1'	4.41 (1H, d, $J = 7.8$ Hz)	104.3	4.35 (1H, d, $J = 7.8$ Hz)	104.3
2'	3.20 (1H, dd, $J = 9.0, 8.0$ Hz)	74.9	3.17 (1H, dd, $J = 9.2, 7.8$ Hz)	74.9
3'	3.39 (1H, t, $J = 9.0$ Hz)	77.8	3.36 (1H, t, $J = 9.1$ Hz)	77.8
4'	3.33 (1H, m)	71.7	3.27 (1H, d, $J = 9.6$ Hz)	71.7
5'	3.55 (1H, m)	75.5	3.49 (1H, m)	75.4
6'	4.32 (1H, dd, $J = 11.9, 6.4$ Hz) 4.53 (1H, dd, $J = 11.9, 2.1$ Hz)	64.5	4.21 (1H, dd, $J = 11.9, 6.6$ Hz) 4.54 (1H, dd, $J = 11.9, 2.1$ Hz)	64.4
1''		168.9		167.9
2''	6.35 (1H, d, $J = 15.9$ Hz)	114.9	5.79 (1H, d, $J = 12.8$ Hz)	116.3
3''	7.64 (1H, d, $J = 15.9$ Hz)	146.9	6.89 (1H, d, $J = 12.8$ Hz)	145.3
4''		127.0		127.6
5'', 9''	7.46 (2H, d, $J = 8.6$ Hz)	131.2 \times 2	7.65 (2H, d, $J = 9.0$ Hz)	133.8 \times 2
6'', 8''	6.81 (2H, d, $J = 8.6$ Hz)	116.9 \times 2	6.77 (2H, d, $J = 9.0$ Hz)	115.9 \times 2
7''		161.4		160.2

SUPPORTING INFORMATION

Table S 2 ^1H (600 MHz) and ^{13}C (150 MHz) data of compounds **3** and **4** (Methanol- d_4).

No.	3		4	
	δ_{H} , mult, J (Hz)	δ_{C}	δ_{H} , mult, J (Hz)	δ_{C}
1		173.7		173.7
2	2.70 (1H, dd, $J = 15.8, 6.9$ Hz) 2.59 (1H, dd, $J = 15.8, 5.8$ Hz)	40.8	2.68 (1H, m) 2.58 (1H, dd, $J = 15.8, 5.7$ Hz)	40.8
3	4.23 (1H, m)	75.7	4.18 (1H, m)	75.8
4	1.84 (1H, m) 1.57 (1H, ddd, $J = 14.0, 5.8, 4.7$ Hz)	45.2	1.82 (1H, m) 1.55 (1H, ddd, $J = 14.1, 5.8, 4.7$ Hz)	45.2
5	3.96 (1H, m)	66.0	3.94 (1H, m)	66.0
6	1.14 (3H, d, $J = 6.2$ Hz)	22.4	1.12 (3H, d, $J = 6.2$ Hz)	23.3
1-OCH ₃	3.67 (3H, s)	52.2	3.68 (3H, s)	52.2
	Glc	Glc	Glc	Glc
1'	4.42 (1H, d, $J = 7.8$ Hz)	103.6	4.38 (1H, d, $J = 7.8$ Hz)	103.7
2'	3.16 (1H, dd, $J = 9.1, 7.8$ Hz)	75.0	3.14 (1H, dd, $J = 9.2, 7.8$ Hz)	75.0
3'	3.38 (1H, t, $J = 9.0$ Hz)	77.8	3.36 (1H, m)	77.8
4'	3.34 (1H, m)	71.8	3.27 (1H, m)	71.7
5'	3.55 (1H, m)	75.4	3.50 (1H, ddd, $J = 9.6, 6.3, 2.1$ Hz)	75.3
6'	4.30 (1H, dd, $J = 11.9, 6.4$ Hz) 4.52 (1H, dd, $J = 11.9, 2.1$ Hz)	64.7	4.23 (1H, dd, $J = 11.9, 6.4$ Hz) 4.48 (1H, dd, $J = 11.9, 2.1$ Hz)	64.5
1''		169.1		168.0
2''	6.36 (1H, d, $J = 16.0$ Hz)	115.0	5.80 (1H, d, $J = 12.8$ Hz)	116.3
3''	7.65 (1H, d, $J = 16.0$ Hz)	146.8	6.88 (1H, d, $J = 12.8$ Hz)	145.4
4''		127.1		127.6
5'', 9''	7.47 (2H, d, $J = 8.6$ Hz)	131.2 × 2	7.67 (2H, d, $J = 8.6$ Hz)	133.8 × 2
6'', 8''	6.91 (2H, d, $J = 8.6$ Hz)	116.8 × 2	6.76 (2H, d, $J = 8.6$ Hz)	115.9 × 2
7''		161.4		160.2

SUPPORTING INFORMATION

D

Table S 3. Experimental $[\alpha]$ data of aglycone of compounds **3** and **4** at 20 °C with 589 nm laser (methanol as the solvent) and the calculated $[\alpha]$ of their aglycone in four absolute configurations.

Comp.	structure	calculated $[\alpha]$	experimental $[\alpha]$
3' a (3R5R)		-27.72718	
3' b (3R5S)		-1.90662	
3' c (3S5R)		4.10722	
3' d (3S5S)		28.43639	
3'			6.6
4'			7.0

SUPPORTING INFORMATION

Table S 4 ^1H (600 MHz) and ^{13}C (150 MHz) data of compounds **5** and **6** (Methanol- d_4).

No.	5		6	
	δ_{H} , mult, J (Hz)	δ_{C}	δ_{H} , mult, J (Hz)	δ_{C}
1		176.4		176.5
2		130.5		130.3
3	7.22 (1H, t, J = 1.8 Hz)	151.7	7.26 (1H, t, J = 1.8 Hz)	151.9
4	5.36 (1H, dt, J = 9.6, 1.8 Hz)	85.8	5.35 (1H, dt, J = 9.0, 1.8 Hz)	86.1
5	2.98 (1H, t, J = 9.6 Hz)	45.7	3.15 (1H, dd, J = 9.6, 9.0 Hz)	44.6
6	5.76 (1H, dd, J = 9.6, 1.8 Hz)	124.4	5.77 (1H, dd, J = 9.6, 1.8 Hz)	124.9
7		139.0		138.3
8	3.96 (2H, s)	68.3	3.95 (2H, s)	68.4
9	1.68 (3H, s)	14.5	1.70 (3H, s)	14.4
10	1.86 (3H, t, J = 1.8 Hz)	10.5	1.86 (3H, t, J = 1.8 Hz)	10.4
1'		137.2		137.1
2'		129.0		129.6
3'	2.05 (1H, dd, J = 16.8, 9.6 Hz) 2.47 (1H, ddd, J = 16.8, 5.4, 1.8 Hz)	41.1	2.24 (1H, dd, J = 16.8, 8.4 Hz) 2.36 (1H, ddd, J = 16.8, 5.4, 1.2 Hz)	42.5
4'	4.11 (1H, m)	72.6	3.97 (1H, m)	73.7
5'	1.46 (1H, t, J = 12.0 Hz) 1.83 (1H, dt, J = 12.0, 3.0 Hz)	47.5	1.54 (1H, t, J = 12.0 Hz) 1.83 (1H, dt, J = 12.0, 2.4 Hz)	45.8
6'		39.5		39.2
7'	1.85 (3H, s)	22.1	1.87 (3H, s)	22.7
8'	0.98 (3H, s)	28.6	0.92 (3H, s)	28.8
9'	0.93 (3H, s)	29.6	1.01 (3H, s)	30.7
		Glc		
1''	4.44 (1H, d, J = 7.8 Hz)	102.4	4.37 (1H, d, J = 7.8 Hz)	102.8
2''	3.16 (1H, dd, J = 9.0, 7.8 Hz)	75.1	3.16 (1H, m)	75.2
3''	3.37 (1H, m)	78.1	3.35 (1H, m)	78.1
4''	3.28 (1H, m)	71.6	3.28 (1H, m)	71.7
5''	3.29 (1H, m)	77.9	3.27 (1H, m)	77.9
6''	3.67 (1H, dd, J = 12.0, 2.4 Hz) 3.88 (1H, dd, J = 12.0, 1.8 Hz)	62.7	3.66 (1H, dd, J = 12.0, 2.4 Hz) 3.86 (1H, dd, J = 12.0, 1.8 Hz)	62.8

SUPPORTING INFORMATION

Table S 5 ^1H (600 MHz) and ^{13}C (150 MHz) data of compound **7** (DMSO- d_6).

No.	δ_{H} , mult, J (Hz)	δ_{C}
1	7.46 (1H, s)	102.5
2		143.5
3		139.2
4		133.0
4a		143.9
5a		156.7
6	6.97 (1H, d, J = 2.0 Hz)	98.3
7		156.9
8	6.77 (1H, dd, J = 8.4, 2.0 Hz)	115.0
9	7.65 (1H, d, J = 8.4 Hz)	120.3
9a		116.0
9b		111.5
4-OCH ₃	3.99 (3H, s)	60.4
	Glc	Glc
1'	4.69 (1H, d, J = 7.6 Hz)	103.8
2'	3.18 (1H, dd, J = 9.0, 7.6 Hz)	73.5
3'	3.36-3.29 (overlapped in DMSO- d_6)	77.4
4'	3.36-3.29 (overlapped in DMSO- d_6)	69.9
5'	3.36-3.29 (overlapped in DMSO- d_6)	76.0
6'	3.51 (1H, dd, J = 11.6, 6.2 Hz) 3.77 (1H, dd, J = 11.6, 1.5 Hz)	60.8

SUPPORTING INFORMATION

Table S 6 ^1H (600 MHz) and ^{13}C (150 MHz) data of compound **8** (Methanol- d_4)

No.	δ_{H} , mult, J (Hz)	δ_{C}
1		131.2
2	7.08 (1H, d, J = 1.8 Hz)	112.7
3		146.6
4		136.6
5		149.2
6	6.94 (1H, d, J = 1.8 Hz)	109.7
1'		129.6
2'		155.2
3'	6.86 (1H, dd, J = 8.0, 1.4 Hz)	117.0
4'	7.10 (1H, td, J = 7.9, 1.6 Hz)	129.1
5'	6.89 (1H, td, J = 8.0, 1.4 Hz)	121.0
6'	7.26 (1H, dd, J = 7.9, 1.6 Hz)	131.5
5-OCH ₃	3.87 (3H, s)	56.8
	Glc	Glc
1''	4.86 (1H, d, J = 7.7 Hz)	104.2
2''	3.38 (1H, m)	74.9
3''	3.55 (1H, m)	78.2
4''	3.44 (1H, m)	71.2
5''	3.47 (1H, m)	77.6
6''	3.72 (1H, dd, J = 12.0, 5.1 Hz) 3.86 (1H, dd, J = 12.0, 2.4 Hz)	62.3

SUPPORTING INFORMATION

Table S 7 ^1H (600 MHz) and ^{13}C (150 MHz) data of compound **9** (Methanol- d_4).

No.	δ_{H} , mult, J (Hz)	δ_{C}
1	5.22 (1H, dd, $J = 17.4, 1.5$ Hz) 4.99 (1H, dd, $J = 10.8, 1.5$ Hz)	111.9
2	6.00 (1H, dd, $J = 17.4, 10.8$ Hz)	145.4
3		84.6
4	1.89 (1H, m) 1.81 (1H, m)	38.6
5	2.01 (1H, m) 1.85 (1H, m)	28.0
6	4.08 (1H, t, $J = 7.2$ Hz)	85.1
7		80.1
8	1.27 (3H, s)	23.7
9	1.23 (3H, s)	22.7
10	1.30 (3H, s)	25.9
	7-Glc	
1'	4.57 (1H, d, $J = 7.8$ Hz)	98.4
2'	3.14 (1H, dd, $J = 9.2, 7.8$ Hz)	75.3
3'	3.37 (1H, t, $J = 9.1$ Hz)	78.0
4'	3.27 (1H, dd, $J = 9.6, 9.1$ Hz)	72.0
5'	3.42 (1H, ddd, $J = 9.5, 6.0, 2.4$ Hz)	76.4
6'	3.99 (1H, dd, $J = 11.0, 6.0$ Hz) 3.56 (1H, dd, $J = 11.4, 2.4$ Hz)	68.2
	6'-Ara	
1''	4.94 (1H, d, $J = 1.2$ Hz)	109.9
2''	3.97 (1H, dd, $J = 3.3, 1.5$ Hz)	83.2
3''	3.82 (1H, dd, $J = 5.9, 3.2$ Hz)	79.0
4''	3.96 (1H, dd, $J = 5.6, 3.4$ Hz)	85.9
5''	3.74 (1H, dd, $J = 11.9, 3.3$ Hz) 3.64 (1H, dd, $J = 11.9, 5.3$ Hz)	63.1

SUPPORTING INFORMATION

Table S 8 ^1H (600 MHz) and ^{13}C -NMR (150 MHz) data of compounds **10** and **11**.

No.	10 (DMSO- d_6)		11 (Methanol- d_4)	
	δ_{H} , mult, J (Hz)	δ_{C}	δ_{H} , mult, J (Hz)	δ_{C}
2		156.5		158.3
3		133.3		134.2
4		176.9		178.9
5		161.1		162.8
6	6.11 (1H, s)	99.4	6.13 (1H, d, J = 2.0 Hz)	98.5
7		165.2		170.1
8	6.30 (1H, s)	93.9	6.30 (1H, d, J = 2.0 Hz)	95.0
9		156.1		157.9
10		102.9		105.0
1'		121.8		123.1
2'	7.51 (1H, d, J = 2.1 Hz)	115.2	7.67 (1H, d, J = 2.1 Hz)	117.0
3'		145.0		145.8
4'		149.0		149.5
5'	6.79 (1H, d, J = 8.5 Hz)	115.7	6.87 (1H, d, J = 8.5 Hz)	115.9
6'	7.64 (1H, dd, J = 8.5 ,2.1 Hz)	120.7	7.56 (1H, dd, J = 8.5 ,2.1 Hz)	122.8
3-O-Gal				
1''	5.28 (1H, d, J = 7.7 Hz)	102.4	5.64 (1H, d, J = 7.7 Hz)	101.0
2''		71.1	3.95 (1H, dd, J = 9.5, 7.9 Hz)	77.3
3''		73.1	3.71 (1H, dd, J = 9.6, 3.5 Hz)	75.5
4''		68.0	3.80 (1H, m)	70.7
5''		73.5	3.65 (1H, t, J = 6.4 Hz)	75.1
6''		65.1	3.75 (1H, dd, J = 10.2, 6.1 Hz) 3.47 (1H, dd, J = 10.2, 6.6 Hz)	66.7
2''-O-Rha				
1			5.21 (1H, d, J = 1.1 Hz)	102.4
2			3.99 (1H, dd, J = 3.3, 1.6 Hz)	72.2
3			3.80 (1H, m)	72.1
4			3.34 (1H, t, J = 10.3 Hz)	73.9
5			4.06 (1H, dd, J = 9.7, 6.2 Hz)	69.7
6			0.97 (3H, d, J = 6.2 Hz)	17.2

SUPPORTING INFORMATION

6''-O-Rha	4.42 (1H, d, $J = 0.7$ Hz)	100.0		
1		70.4	4.55 (1H, d, $J = 1.4$ Hz)	101.7
2		70.6	3.58 (1H, dd, $J = 3.4, 1.6$ Hz)	71.9
3		71.9	3.51 (1H, dd, $J = 9.5, 3.5$ Hz)	72.1
4		68.3	3.28 (1H, t, $J = 9.5$ Hz)	73.7
5	1.07 (3H, d, $J = 6.2$ Hz)	17.9	3.54 (1H, dd, $J = 6.2, 3.3$ Hz)	69.5
6	4.42 (1H, d, $J = 0.7$ Hz)	100.0	1.19 (3H, d, $J = 6.2$ Hz)	17.8

SUPPORTING INFORMATION

Table S 9 ^{13}C -NMR (150 MHz) data of compounds **12-15** (DMSO- d_6)

No.	12	13	14	15
2	156.4	156.4	156.2	156.2
3	133.4	133.2	133.2	133.2
4	177.2	177.3	177.4	177.4
5	161.2	161.2	161.2	161.2
6	99.1	99.0	99.0	98.9
7	164.1	165.4	164.7	164.8
8	93.7	93.7	93.8	93.8
9	156.0	156.0	156.5	156.5
10	103.3	103.5	103.5	103.8
1'	120.9	121.6	120.9	120.9
2'	115.2	115.2	131.0	130.9
3'	144.9	144.9	115.1	115.1
4'	148.9	148.7	160.4	160.0
5'	115.8	116.1	115.1	115.1
6'	122.0	121.1	131.0	130.9
	3-O-Gal	3-O-Glc	3-O-Gal	3-O-Glc
1''	102.0	101.0	101.8	100.9
2''	71.2	74.1	71.2	74.2
3''	73.2	77.6	73.1	77.5
4''	67.9	69.9	67.9	70.0
5''	75.8	76.5	75.8	76.5
6''	60.1	61.0	60.2	61.0

SUPPORTING INFORMATION

Table S 10 ^{13}C -NMR (150 MHz) data of compounds **16-21** (DMSO- d_6)

No.	16	17	18	19	20	21
2	164.0	163.9	163.6	164.0	163.7	164.0
3	102.6	102.4	103.1	102.4	102.6	102.4
4	182.0	182.1	182.1	182.0	182.1	182.1
5	161.2	161.2	161.1	161.2	161.2	161.3
6	98.2	98.2	109.5	97.9	98.2	98.4
7	162.8	162.9	163.6	162.5	162.8	162.7
8	103.7	104.7	93.2	102.4	104.0	104.5
9	156.0	156.0	156.9	156.5	156.0	155.9
10	103.8	104.0	103.1	103.8	104.2	104.1
1'	121.7	121.6	121.6	121.6	121.6	121.6
2', 6'	129.0 \times 2	129.0 \times 2	128.8 \times 2	129.0 \times 2	128.7 \times 2	129.0 \times 2
3', 5'	115.9 \times 2	115.9 \times 2	116.5 \times 2	115.9 \times 2	115.9 \times 2	115.9 \times 2
4'	160.6	160.4	161.7	160.8	160.6	160.7
	8-Glc	8-Glc	6-Glc	8-Glc	8-Glc	8-Glc
1''	73.5	73.4	73.7	70.9	73.7	71.7
2''	68.5	70.9	70.3	72.5	70.7	75.1
3''	79.8	78.7	79.5	75.7	78.2	79.9
4''	68.1	70.6	70.6	70.5	70.3	70.7
5''	81.5	81.9	82.0	80.0	78.3	81.8
6''	60.6	61.3	61.9	61.0	64.1	61.2
	3''-O-COCH ₃			2''-O-COCH ₃	6''-O-COCH ₃	2''-O-Rha
1'''	169.9			169.2	170.4	100.3
2'''	21.2			20.5	20.6	70.3
3'''						71.5
4'''						70.5
5'''						68.3
6'''						17.2

SUPPORTING INFORMATION

Table S 11 Antioxidant activity of separation fractions of the leaves of *Crataegus pinnatifida* Bge. in vitro

Fraction	IC ₅₀ (mg/ml)		No.	IC ₅₀ (mg/ml)	
	DPPH•	ABTS•+		DPPH•	ABTS•+
Fr.1	30.9±0.94	5.80±0.03	Fr.3.5	>80	29.2±0.81
Fr.2	52.4±1.14	22.2±0.44	Fr.5.1	>80	>80
Fr.3	49.1±1.20	13.3±0.21	Fr.5.2	19.9±1.25	11.1±0.42
Fr.4	>80	27.6±0.80	Fr.5.3	13.1±0.87	13.6±0.46
Fr.5	21.1±1.07	4.9±0.09	Fr.5.4	51.8±1.53	26.4±0.73
Fr.6	>80	65.7±1.12	VC	33.7±1.41	6.4±0.13

(VC: Positive control)

SUPPORTING INFORMATION

Table S 12 Antioxidant activity of compounds **1-21** in vitro

No.	IC ₅₀ (μM)		No.	IC ₅₀ (μM)	
	DPPH•	ABTS ^{·+}		DPPH•	ABTS ^{·+}
1	>200	63.1±0.01	12	125.2±0.64	13.0±0.86
2	>200	>200	13	>200	28.6±1.04
3	>200	69.1±0.99	14	>200	143.7±1.12
4	>200	>200	15	>200	155.5±0.86
5	>200	>200	16	>200	>200
6	>200	>200	17	>200	164.2±1.30
7	149.3±0.65	16.5±1.42	18	>200	38.2±1.07
8	54.9±0.95	11.0±0.03	19	>200	>200
9	>200	165.5±1.44	20	>200	>200
10	83.9±0.60	3.4±0.28	21	>200	118.3±0.96
11	>200	59.5±0.75	VC	157.5±0.90	17.1±1.05

(VC: Positive control)

SUPPORTING INFORMATION

Table S 13 The protective effect of flavonoids **1-21** on SH-SY5Y cells at 25, 50, 100 μ M.

No.	concentration (μ M)				No.	concentration (μ M)			
	25	50	100	300		25	50	100	300
1	48.2	47.7	53.0		13	53.7	67.0	81.9	
2	37.7	40.6	37.4		VE				75.1
3	48.2	46.6	40.1		14	57.5	76.1	76.1	
4	48.0	47.0	40.6		15	65.5	68.7	70.6	
VE			55.1		VE				71.5
5	49.3	50.9	57.0		16	48.2	54.4	75.4	
6	49.6	50.8	46.6		17	49.3	68.9	70.6	
7	53.5	45.3	52.2		VE				76.6
8	46.4	52.5	43.1		18	59.0	73.3	86.7	
9	56.6	62.0	58.5		19	66.4	68.2	83.2	
VE			72.3		VE				58.7
10	73.3	71.5	78.1		20	50.2	55.5	64.7	
11			66.0		21	42.1	50.4	56.5	
VE			72.9		VE				59.0
12	61.5	72.7	79.3						

(Positive control: VE)

SUPPORTING INFORMATION

Table S 14 The protective effect of flavonoids **10**, **12-21** on alcoholic hepatocyte injury at 25, 50, 100 μ M.

No.	concentration (μ M)			No.	concentration (μ M)		
	25	50	100		25	50	100
10	59.9	61.5	46.3	17	58.8	77.7	58.6
12	59.8	59.9	45.3	18	62.4	72.5	59.3
13	71.8	81.7	73.5	19	56.4	73.1	58.9
14	65.1	67.0	71.6	20	64.8	78.0	74.3
15	62.1	73.4	67.4	21	66.6	62.0	65.6
16	66.0	78.0	69.0	3'g	85.2	99.8	92.3

(3'g: Positive control)