

## SUPPLEMENTARY MATERIAL

# Two new alkaloids and a new butenolide derivative from the Beibu Gulf sponge-derived fungus *Penicillium* sp. SCSIO 41413

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**Abstract:** Marine sponge-derived fungi have been proven to be a prolific source of bioactive natural products. Two new alkaloids, polonimides E (**1**) and D (**2**), and a new butenolide derivative, eutypoid F (**11**), were isolated from the Beibu Gulf sponge-derived fungus, *Penicillium* sp. SCSIO 41413, together with thirteen known compounds (**3–10**, **12–16**). Their structures were determined by detailed NMR, MS spectroscopic analyses, and electronic circular dichroism (ECD) analyses. Butenolide derivatives **11** and **12** exhibited inhibitory effect against the enzyme PI3K with IC<sub>50</sub> values of 1.7  $\mu$ M and 9.8  $\mu$ M, respectively. The molecular docking was also performed to understand the inhibitory activity, while **11** and **12** showed obvious protein/ligand-binding effects to the PI3K protein. Moreover, **4** and **15** displayed obvious inhibitory activity against LPS-induced NF- $\kappa$ B activation in RAW264.7 cells at 10  $\mu$ M.

**Keywords:** Sponge-derived fungus; *Penicillium* sp.; PI3K; NF- $\kappa$ B

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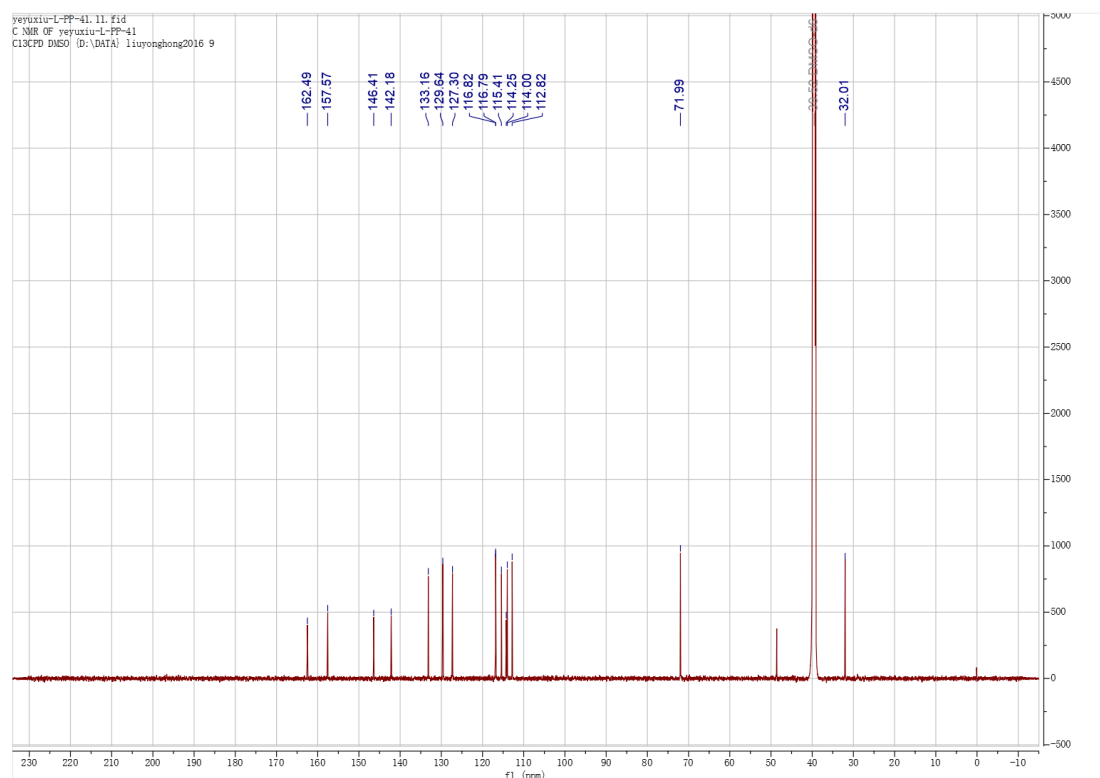
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The physicochemical data of the known compounds 3~10 and 12~16

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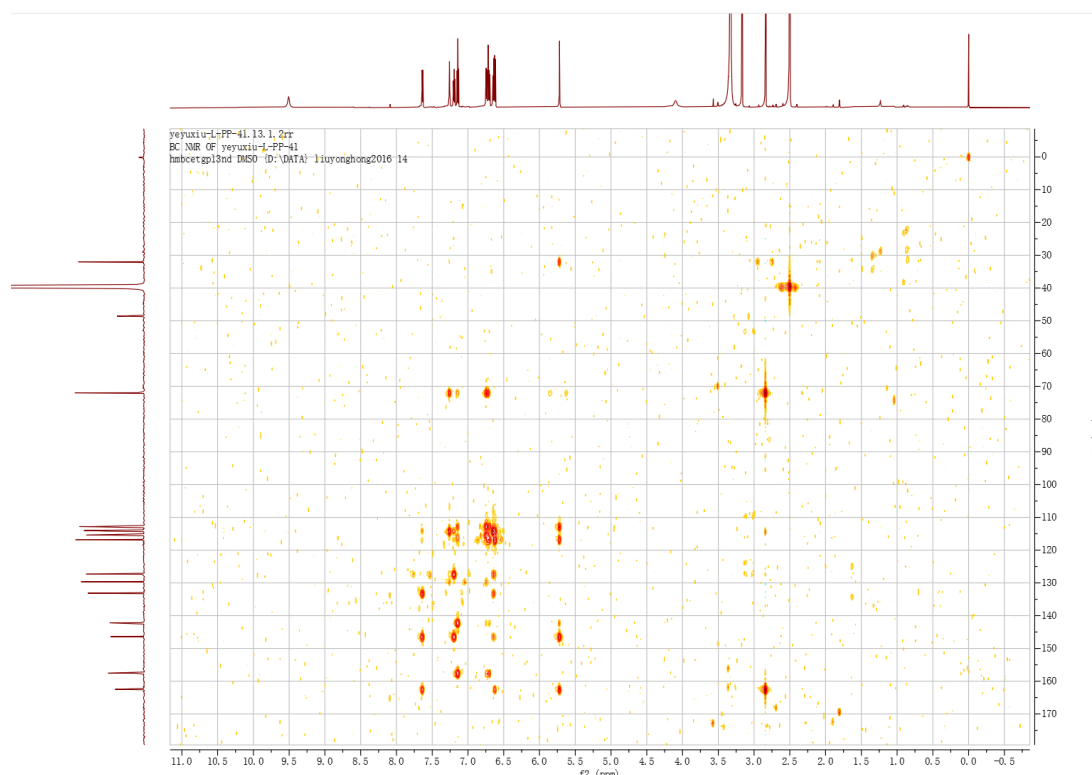
**Figure S1. The  $^1\text{H}$  NMR spectrum of compound 1 in DMSO.**



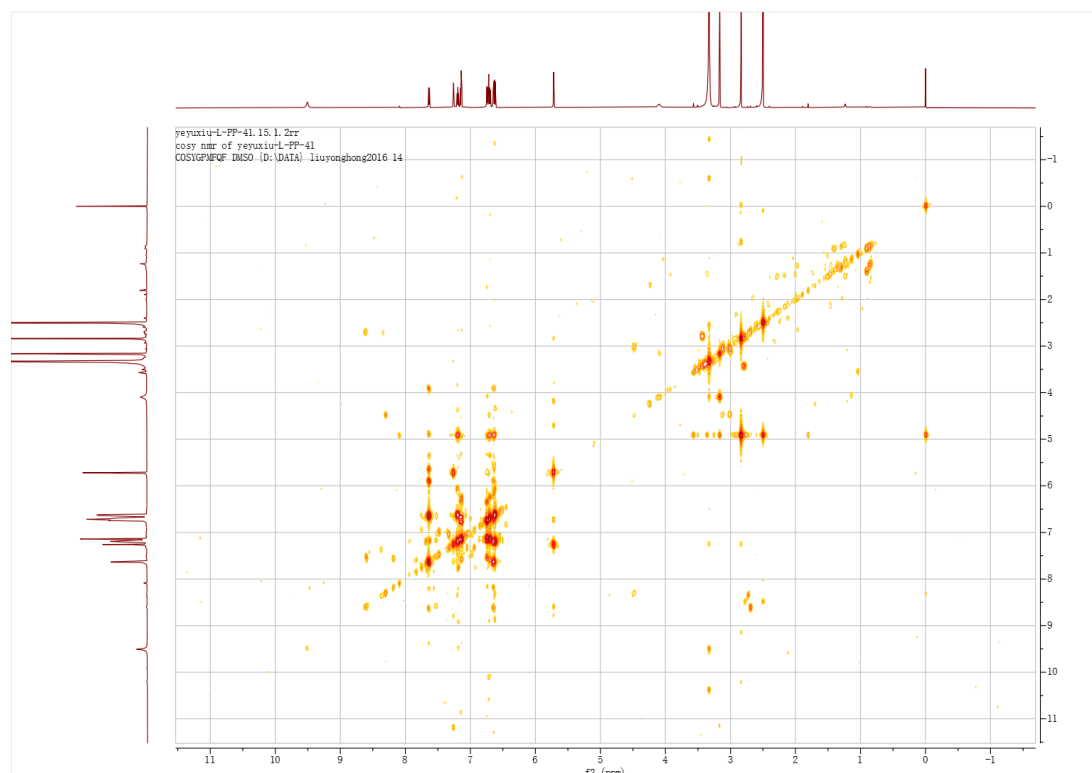
**Figure S2. The  $^{13}\text{C}$  NMR spectrum of compound 1 in DMSO.**



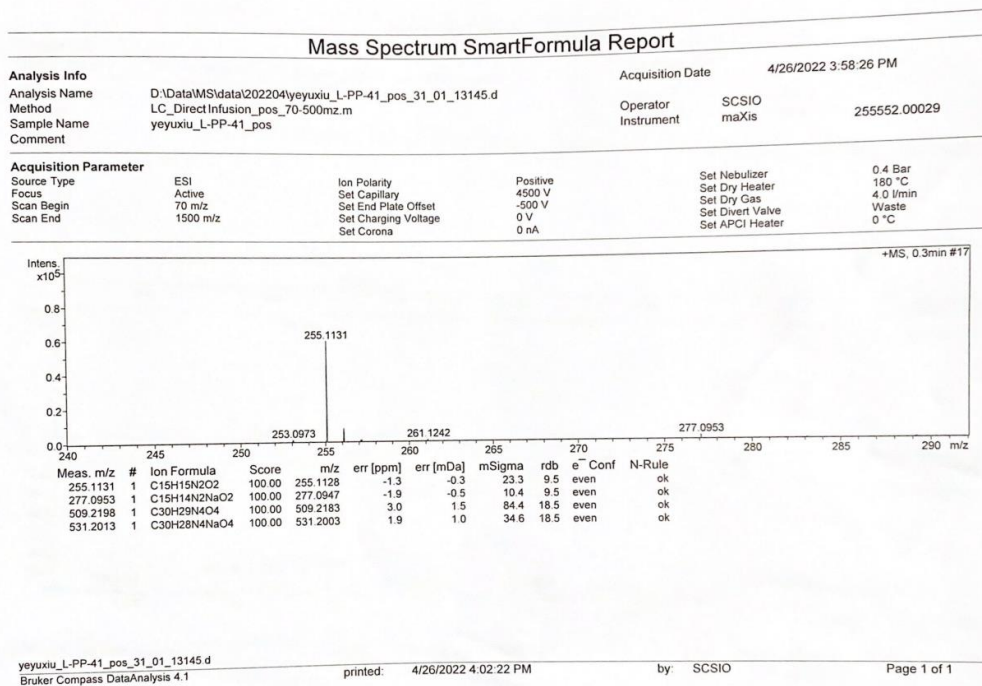
**Figure S3. The HSQC spectrum of compound 1 in DMSO.**



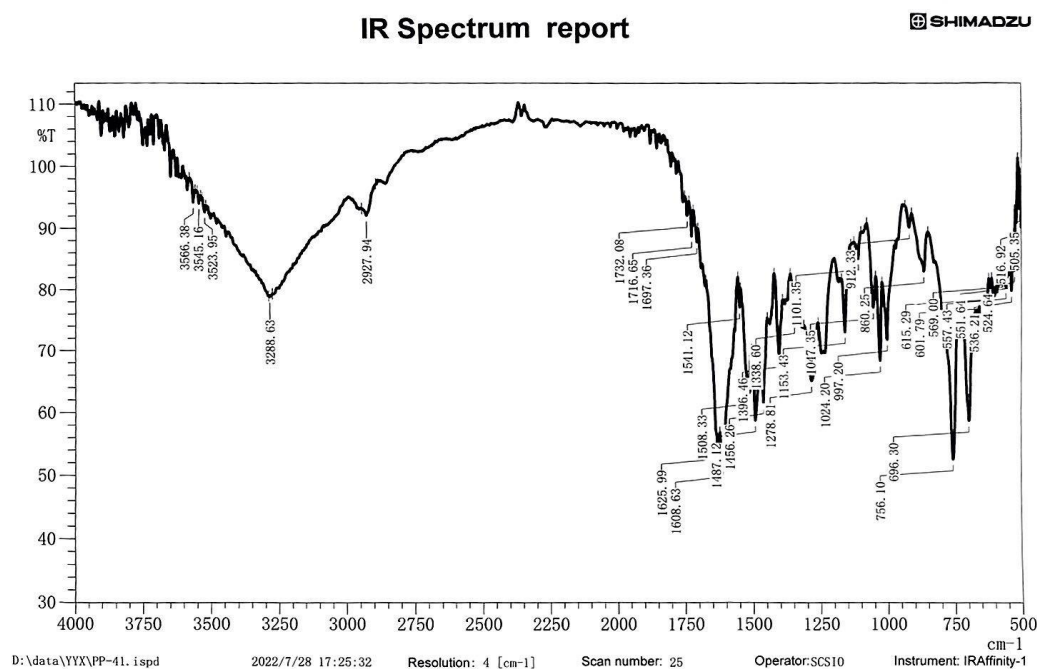
**Figure S4. The HMBC spectrum of compound 1 in DMSO.**



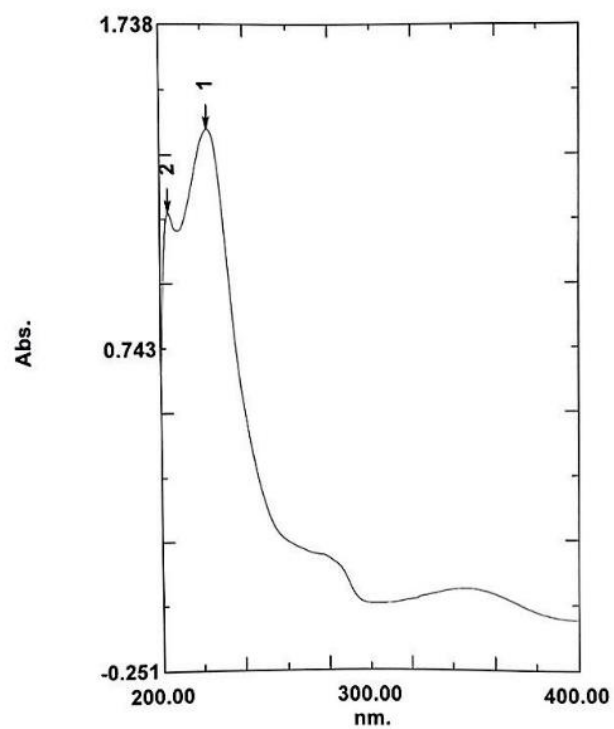
**Figure S5. The  $^1\text{H}$ - $^1\text{H}$  COSY spectrum of compound 1 in DMSO.**



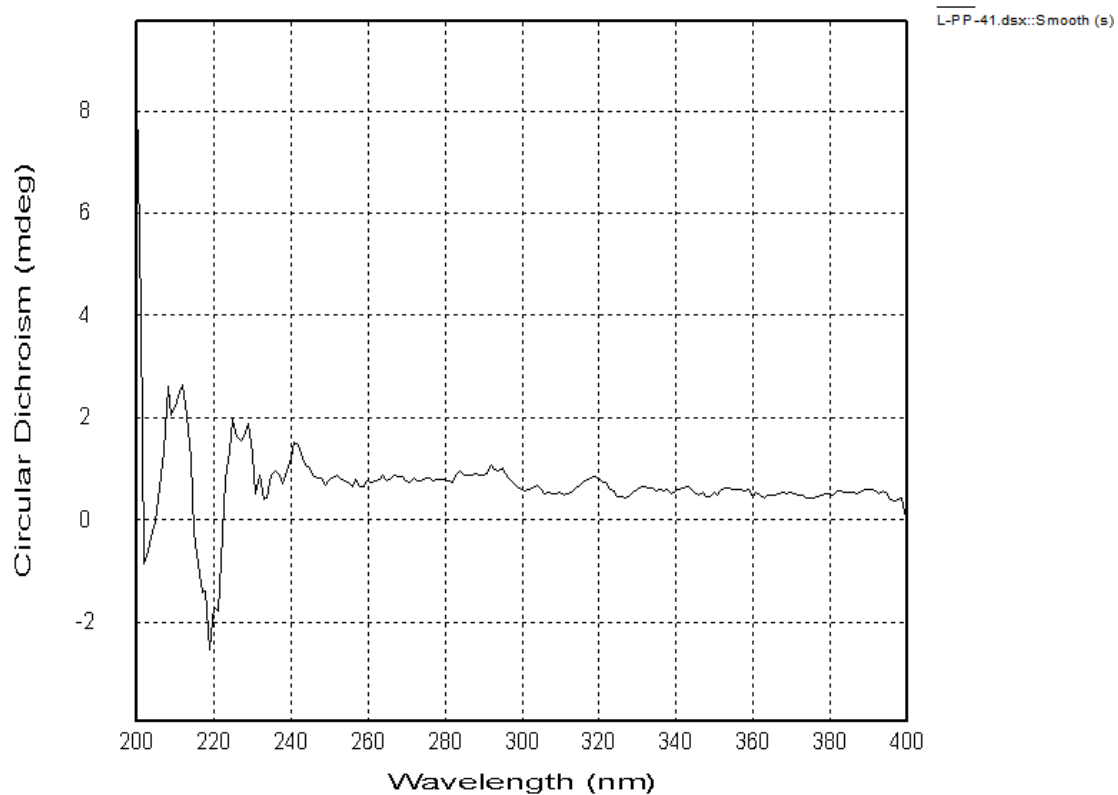
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**Figure S7. The IR spectrum of compound 1 in DMSO.**



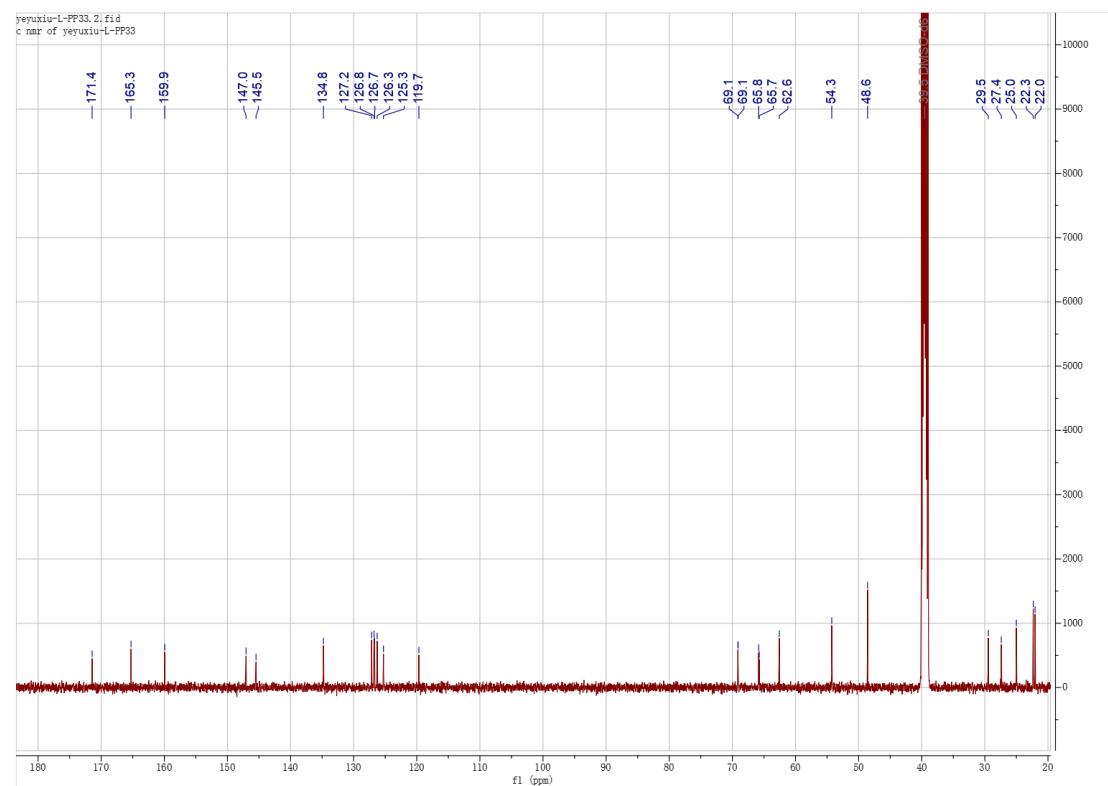
**Figure S8. The UV spectrum of compound 1 in DMSO.**



**Figure S9. The experimental ECD spectrum of compound 1.**

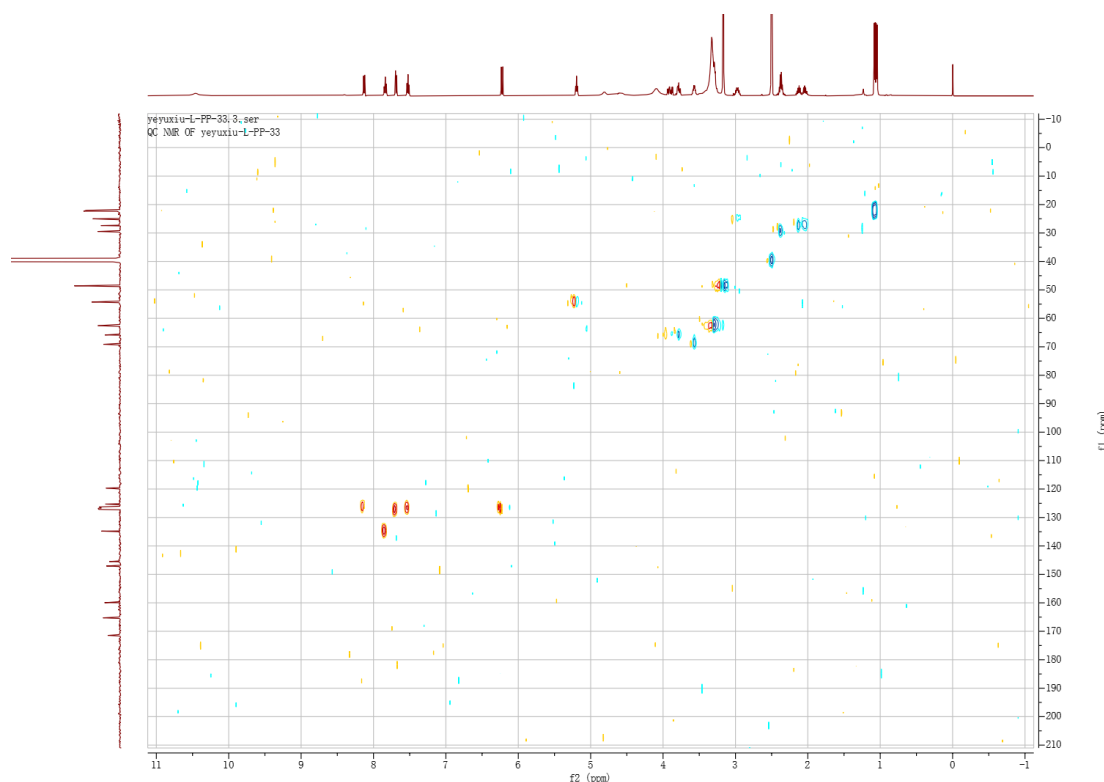


**Figure S10. The  $^1\text{H}$  NMR spectrum of compound 2 in DMSO.**

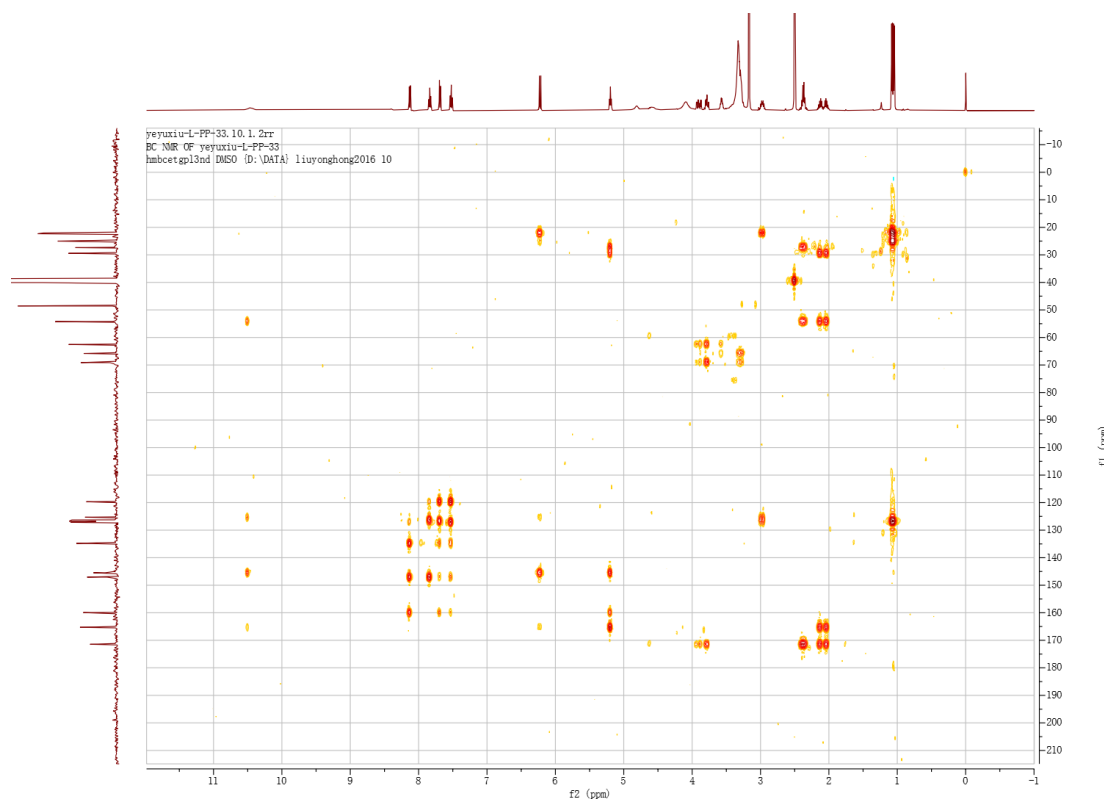


**Figure S11. The  $^{13}\text{C}$  NMR spectrum of compound 2 in DMSO.**

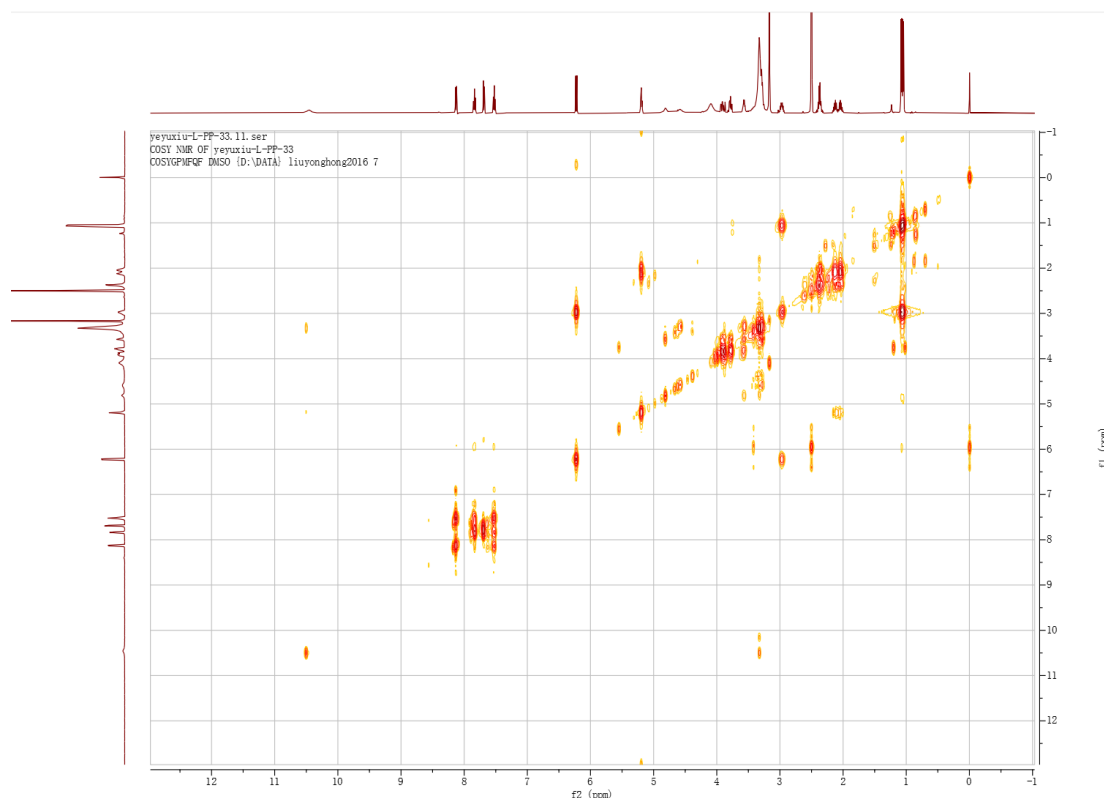




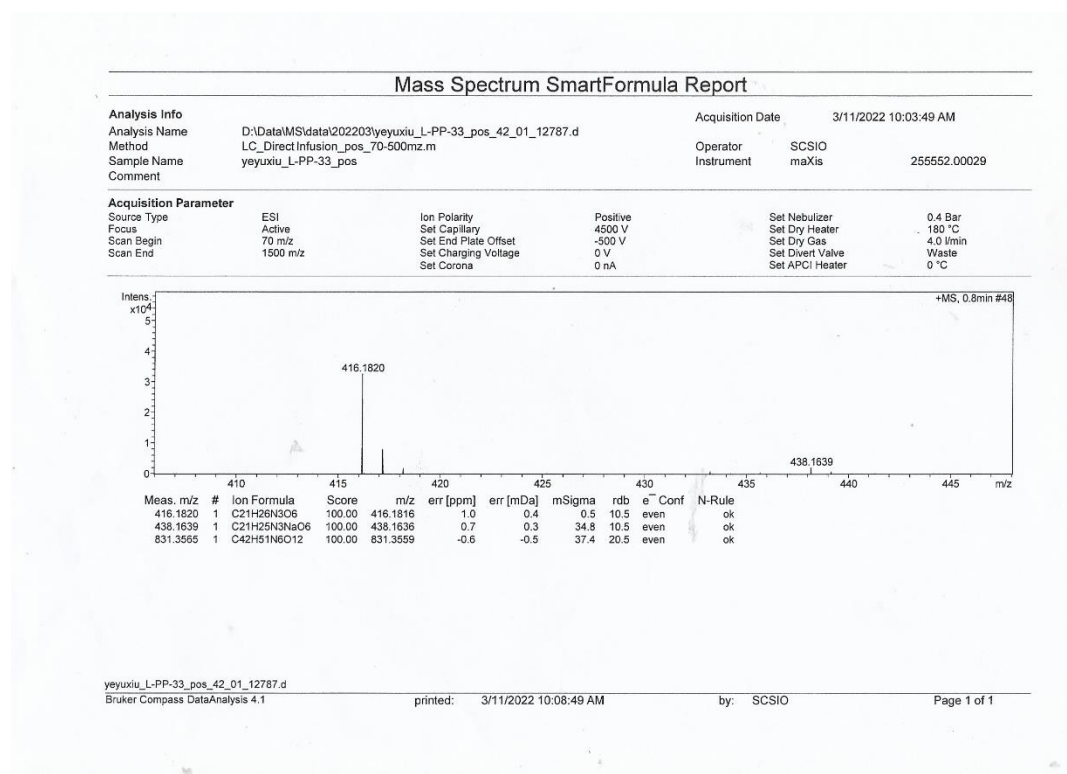
**Figure S12. The HSQC spectrum of compound 2 in DMSO.**



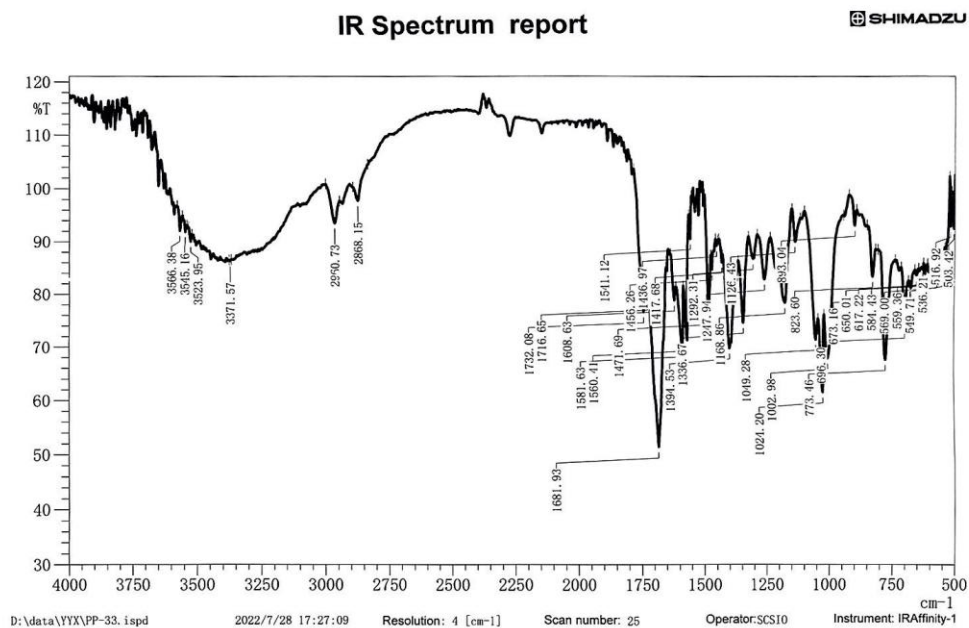
**Figure S13. The HMBC spectrum of compound 2 in DMSO.**



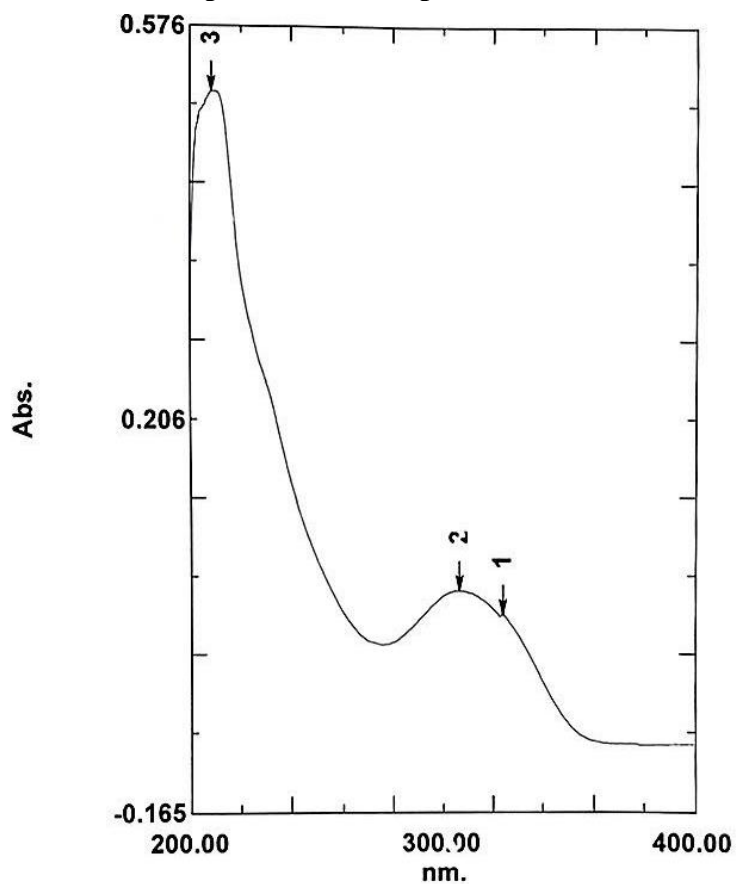
**Figure S14.** The  $^1\text{H}$ - $^1\text{H}$  COSY spectrum of compound 2 in DMSO.



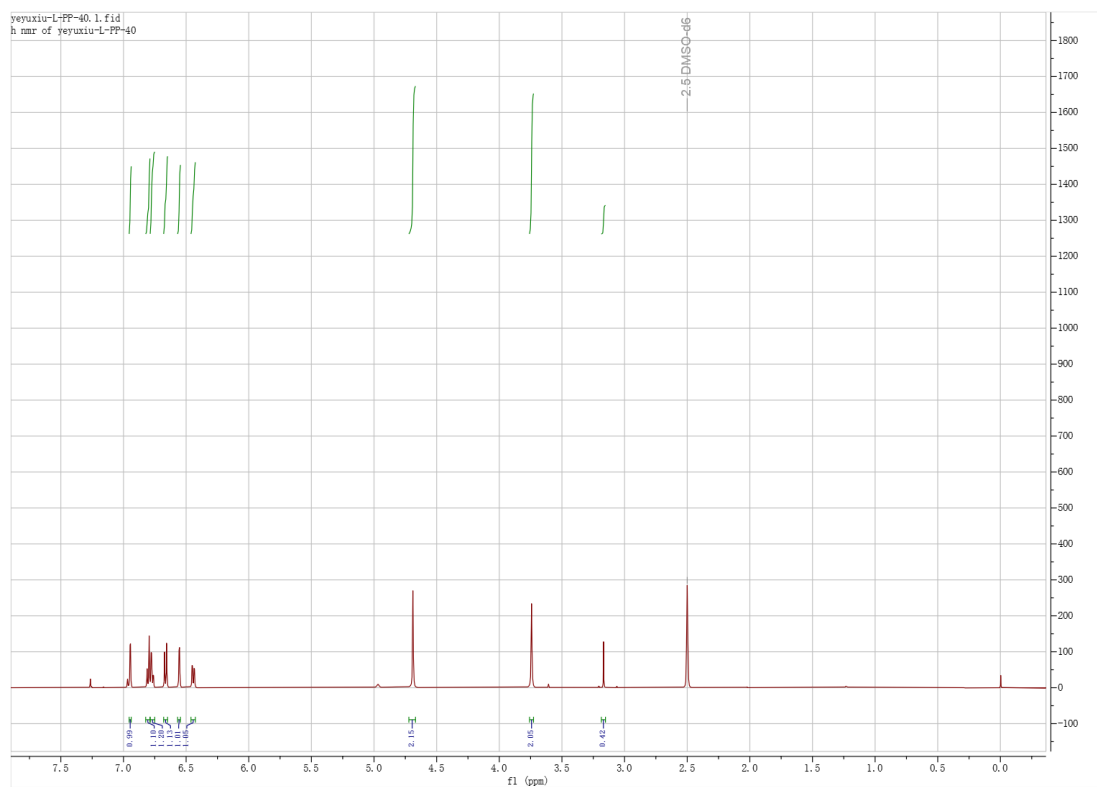
**Figure S15.** The HRESIMS spectrum of compound 2 in DMSO.



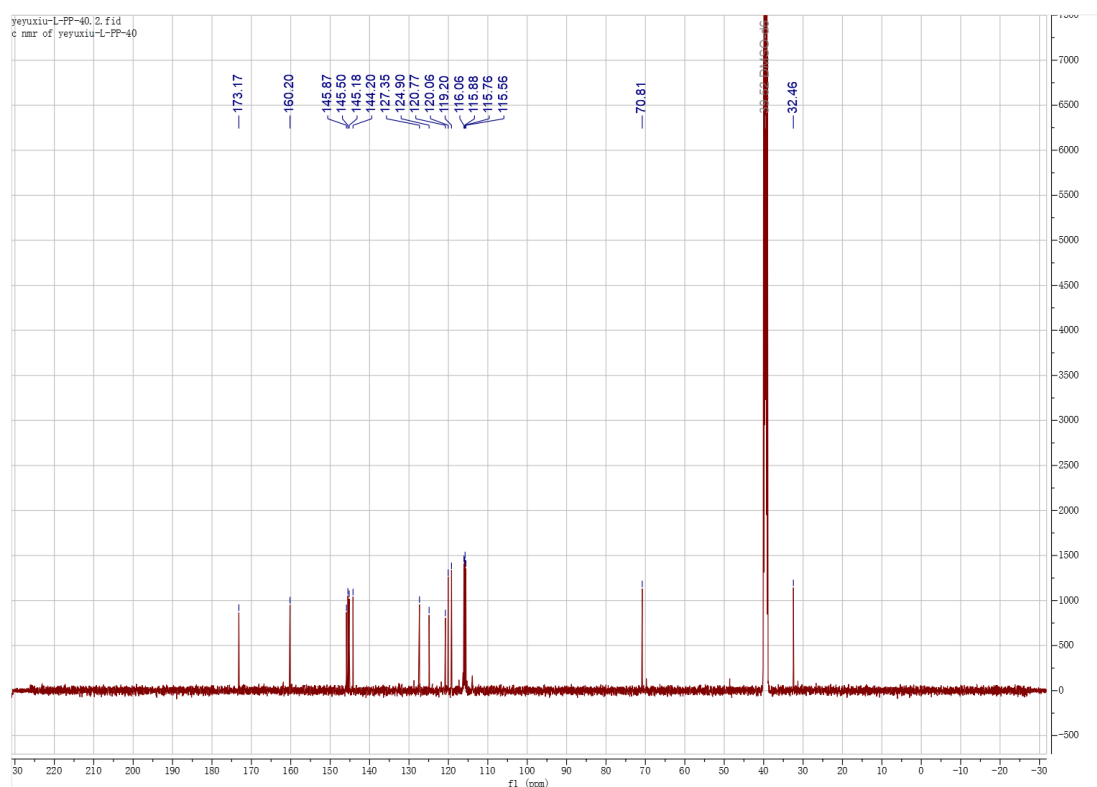
**Figure S16. The IR spectrum of compound 2 in DMSO.**



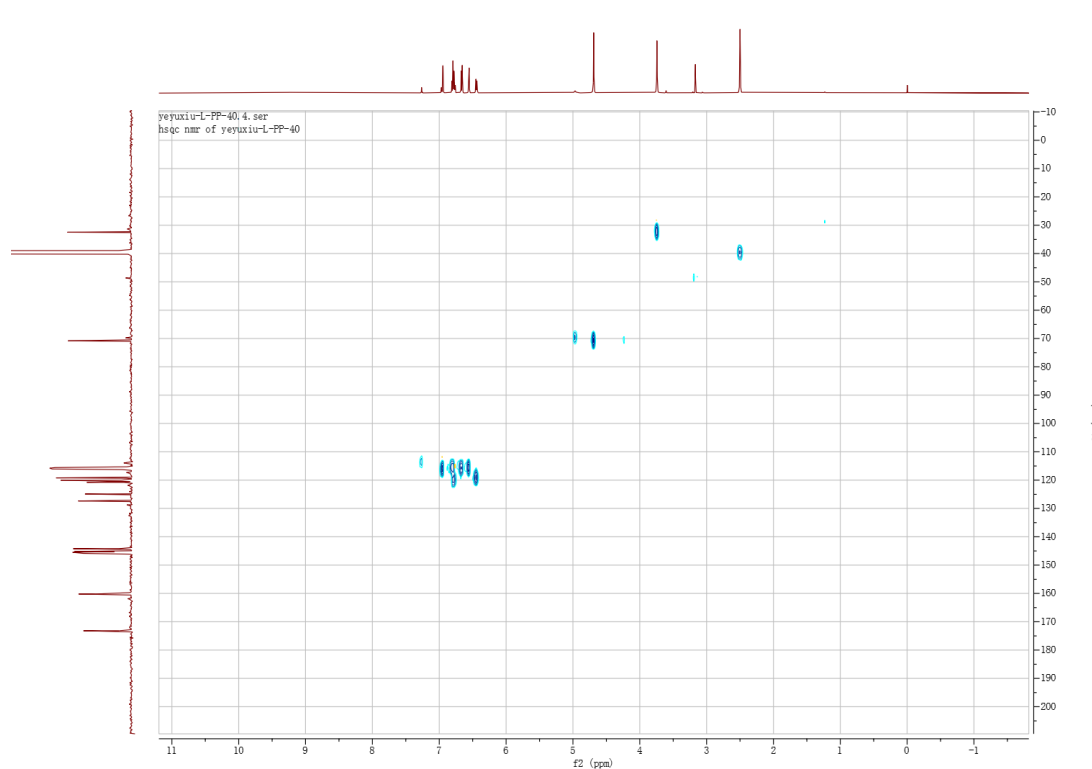
**Figure S17. The UV spectrum of compound 2 in DMSO.**



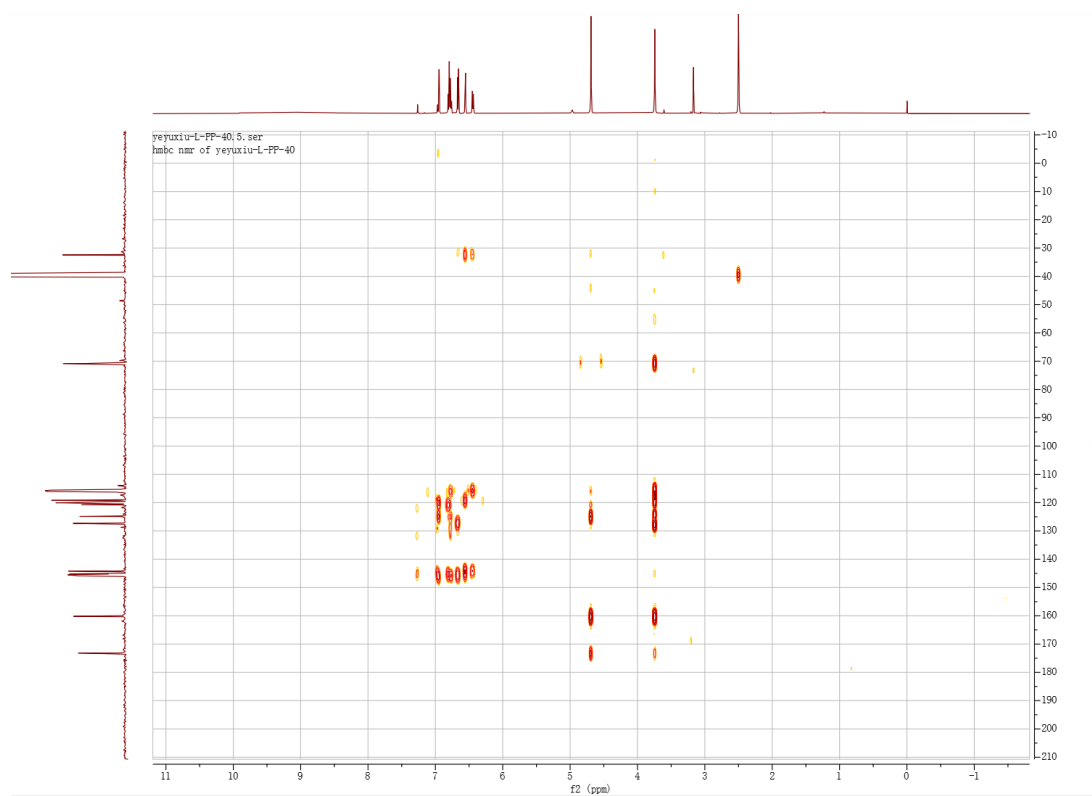
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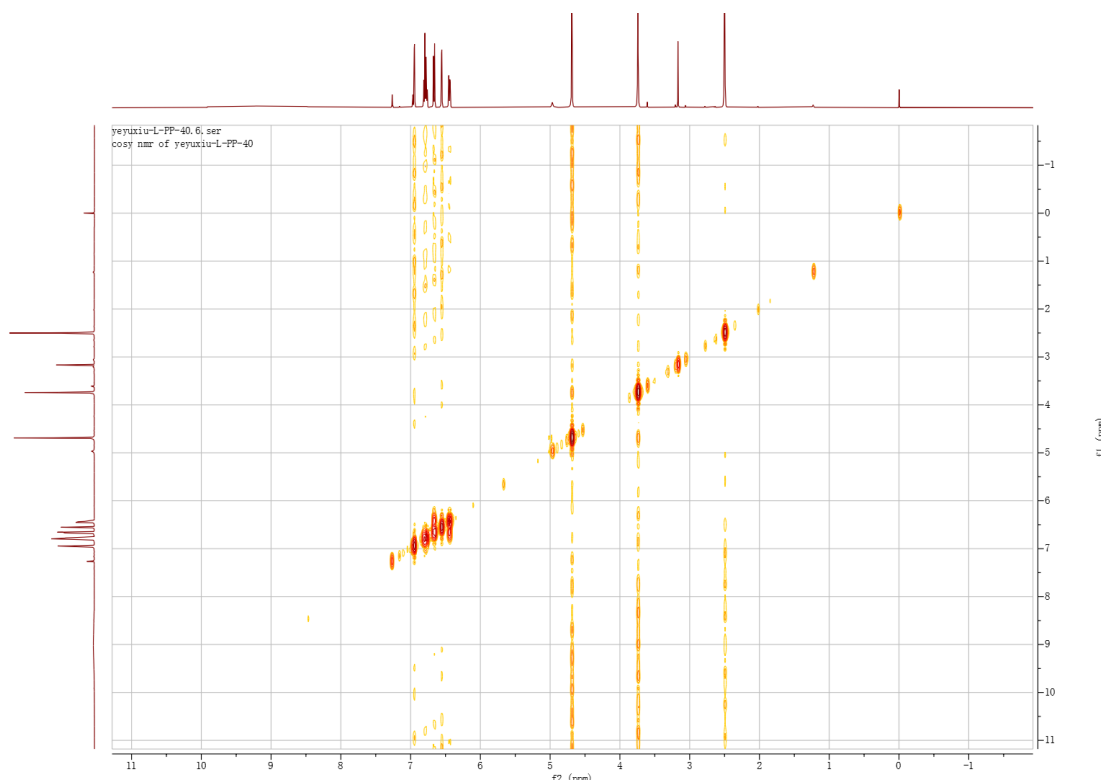
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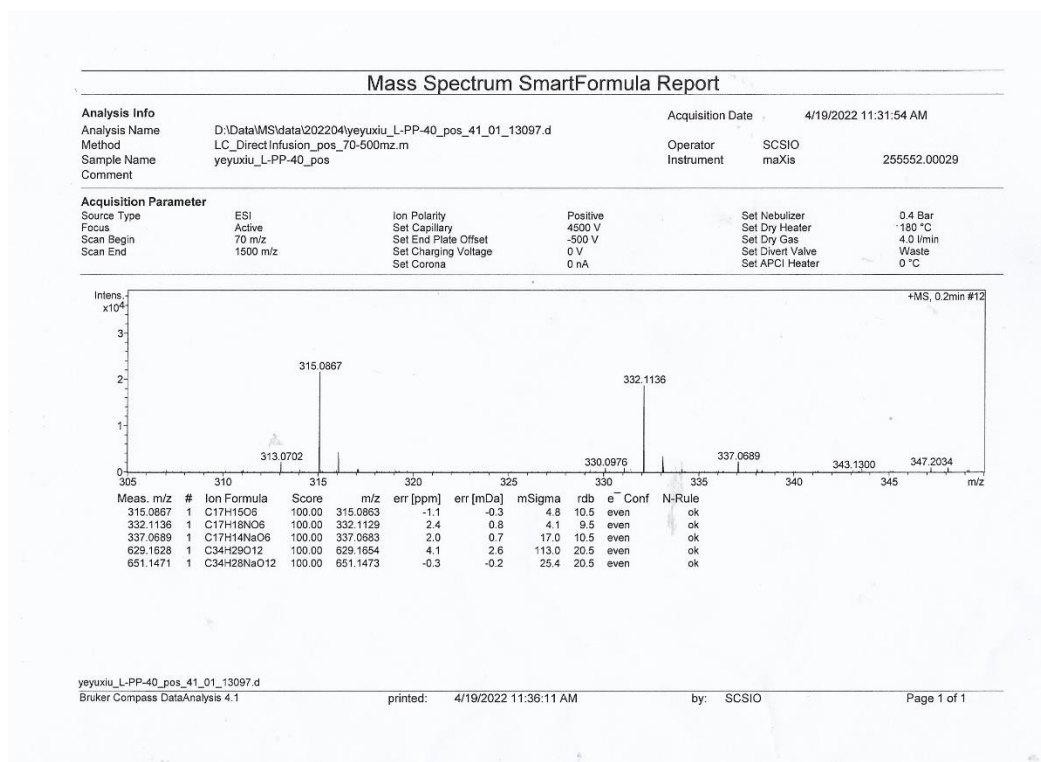
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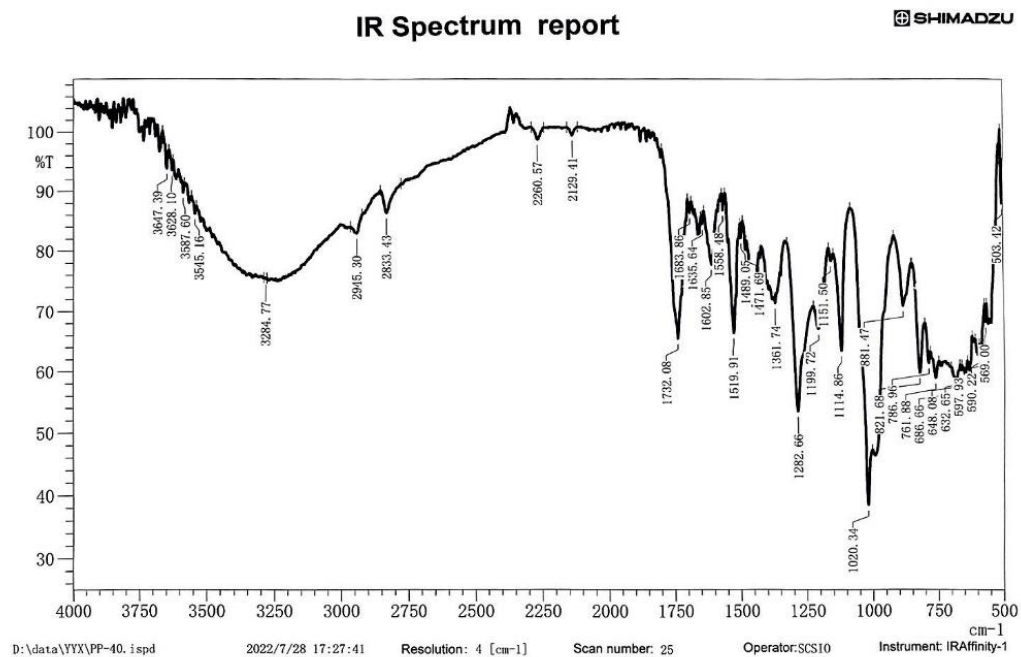
**Figure S21. The HMBC spectrum of compound 11 in DMSO.**



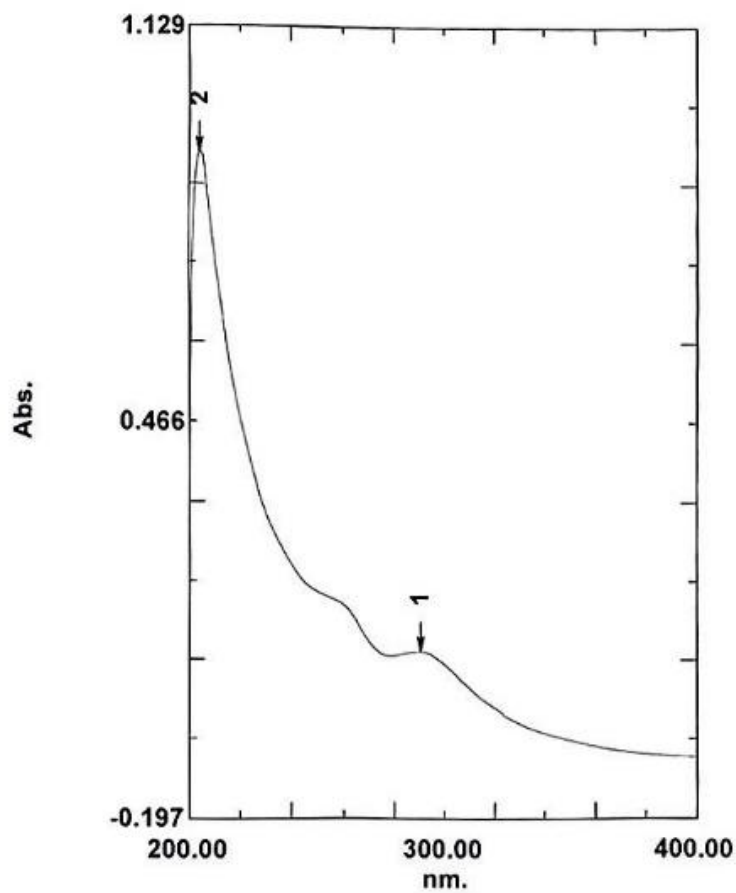
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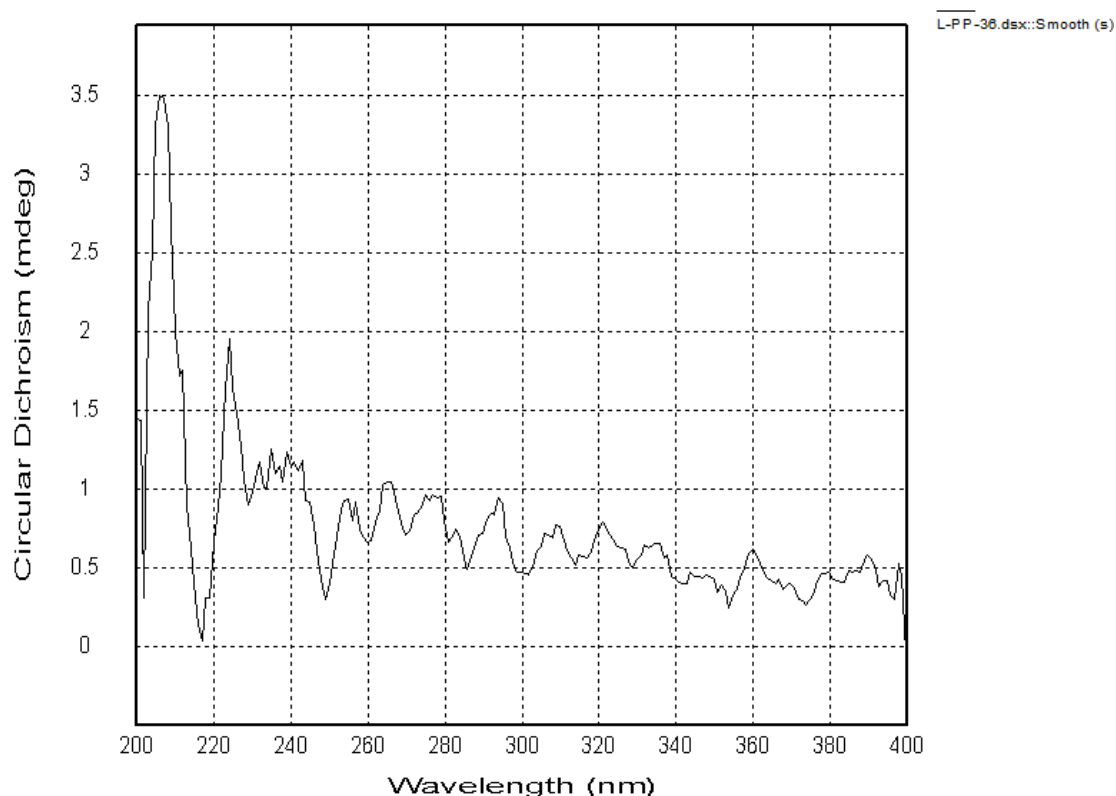
**Figure S23.** The HRMS spectrum of compound 11 in DMSO.



**Figure S24.** The IR spectrum of compound 11 in DMSO.



**Figure S25.** The UV spectrum of compound 11 in DMSO.



**Figure S26. The experimental ECD spectrum of compound 12 in DMSO.**

### The physicochemical data of the known compounds 3~10 and 12~16

polonimide B (**3**): white oil;  $[\alpha]_D^{25}$  -5.42 (c 0.04, MeOH),  $^1\text{H}$  NMR (500 MHz,  $\text{DMSO}-d_6$ )  $\delta_{\text{H}}$  10.49 (1H, s, H-2), 8.16 (1H, dd,  $J = 8.0, 1.5$  Hz, H-10), 7.86 (1H, ddd,  $J = 8.5, 7.1, 1.6$  Hz, H-8), 7.65 (1H, d,  $J = 8.1$  Hz, H-7), 7.57 (1H, t,  $J = 7.6$  Hz, H-9), 7.26 (1H, s, 17-NH<sub>2</sub>), 6.71 (1H, s, 17-NH<sub>2</sub>), 5.54 (1H, d,  $J = 9.8$  Hz, H-18), 5.12 (1H, t,  $J = 6.6$  Hz, H-14), 3.76 (1H, d,  $J = 9.8, 6.6$  Hz, H-19), 2.18-2.07 (1H, m, H-16), 1.20 (3H, d,  $J = 6.6$  Hz, H-21), 1.02 (3H, d,  $J = 6.6$  Hz, H-20).  $^{13}\text{C}$  NMR (126 MHz,  $\text{DMSO}-d_6$ )  $\delta_{\text{C}}$  172.5 (C-17, C), 165.5 (C-1, C), 159.7 (C-12, C), 146.8 (C-6, C), 144.8 (C-4, C), 134.7 (C-8, CH), 131.0 (C-18, CH), 127.5 (C-7, CH), 127.3 (C-9, CH), 126.3 (C-10, CH), 124.8 (C-3, C), 119.8 (C-11, C), 54.7 (C-14, C), 30.9 (C-16, CH<sub>2</sub>), 28.0 (C-15, CH<sub>2</sub>), 26.5 (C-19, CH), 23.2 (C-20, CH<sub>3</sub>), 22.4 (C-21, CH<sub>3</sub>).

Polonimide A (**4**): yellow oil;  $[\alpha]_D^{25}$  -33.37 (c 0.1, MeOH),  $^1\text{H}$  NMR (500 MHz,  $\text{CDCl}_3$ )  $\delta_{\text{H}}$  8.26 (1H, dd,  $J = 8.1, 1.5$  Hz, H-10), 7.76 (1H, ddd,  $J = 8.5, 7.0, 1.5$  Hz, H-8), 7.69 (1H, d,  $J = 7.7$  Hz, H-7), 7.50-7.42 (1H, m, H-9), 6.49 (1H, d,  $J = 10.1$  Hz, H-18), 5.51 (1H, t,  $J = 6.8$  Hz, H-14), 3.57 (3H, s, 17-OCH<sub>3</sub>), 2.83 (1H, q,  $J = 10.0, 8.2$  Hz, H-19), 2.54 (2H, dd,  $J = 8.7, 6.7$  Hz, H-16),



2.31-2.26 (2H, m, H-15), 1.21 (3H, d,  $J = 6.5$  Hz, H-21), 1.19 (3H, d,  $J = 6.4$  Hz, H-20).  $^{13}\text{C}$  NMR (126 MHz,  $\text{CDCl}_3$ )  $\delta_{\text{C}}$  172.4 (C-17, C), 166.5 (C-1, C), 160.7 (C-12, C), 147.5 (C-6, C), 144.6 (C-4, C), 134.9 (C-8, CH), 128.0 (C-18, CH), 127.7 (C-9, CH), 127.1 (C-3, C), 127.0 (C-7, CH), 124.7 (C-10, CH), 120.2 (C-11, C), 54.7 (C-14, CH), 51.9 (17-OCH<sub>3</sub>, CH<sub>3</sub>), 30.1 (C-16, CH<sub>2</sub>), 28.5 (C-15, CH<sub>2</sub>), 26.3 (C-19, CH), 22.5 (C-21, CH<sub>3</sub>), 22.4 (C-20, CH<sub>3</sub>).

aurantioid C (**5**): white powder;  $[\alpha]_{\text{D}}^{25}$  -16.6 (c 0.1, MeOH),  $^1\text{H}$  NMR (500 MHz,  $\text{CDCl}_3$ )  $\delta_{\text{H}}$  8.25 (1H, d,  $J = 8.0$  Hz, H-10), 7.78 (2H, d,  $J = 7.6$  Hz, H-7, H-8), 7.49 (1H, t,  $J = 7.1$  Hz, H-9), 6.53 (1H, d,  $J = 9.9$  Hz, H-18), 5.49 (1H, s, H-14), 2.70 (1H, s, H-19), 2.47 (2H, s, H-16), 2.24 (2H, s, H-15), 1.21 (6H, d,  $J = 6.2$  Hz, H-20, H-21).  $^{13}\text{C}$  NMR (126 MHz,  $\text{CDCl}_3$ )  $\delta_{\text{C}}$  174.2 (C-17, C), 166.6 (C-1, C), 160.9 (C-12, C), 147.5 (C-6, C), 144.8 (C-4, C), 134.9 (C-9, C), 128.4 (C-18, CH), 128.4 (C-7, CH), 127.6 (C-9, CH), 127.0 (C-10, CH), 126.9 (C-3, C), 120.0 (C-11, C), 54.8 (C-14, CH), 31.8 (C-16, CH<sub>2</sub>), 29.0 (C-15, CH<sub>2</sub>), 26.2 (C-19, CH), 22.5 (C-20, CH<sub>3</sub>), 22.4 (C-21, CH<sub>3</sub>).

fructigenine A (**6**): white powder;  $[\alpha]_{\text{D}}^{25}$  -31.4 (c 0.1, MeOH),  $^1\text{H}$  NMR (500 MHz,  $\text{DMSO}-d_6$ )  $\delta_{\text{H}}$  8.20 (1H, s, 2-NH), 7.84 (1H, d,  $J = 8.1$  Hz, H-7), 7.38 (1H, dd,  $J = 7.7, 1.3$  Hz, H-8), 7.31 – 7.22 (6H, m, H-10, 14, 15, 16, 17, 18), 7.13 (1H, d,  $J = 7.5, 1.1$  Hz, H-9), 5.92 (1H, s, H-5a), 5.64 (1H, dd,  $J = 17.3, 10.8$  Hz, H-20), 5.06-5.01 (2H, m, H-21), 4.44 (1H, t,  $J = 4.6$  Hz, H-13), 3.60-3.53 (1H, m, H-11), 3.12 (1H, dd,  $J = 14.0, 4.3$  Hz, Ha-12), 3.01 (1H, dd,  $J = 14.1, 5.1$  Hz, Hb-12), 2.55 (3H, s, H<sub>3</sub>-23), 2.32 (1H, dd,  $J = 12.3, 5.3$  Hz, Ha-11), 1.61 (1H, t,  $J = 12.0$  Hz, Hb-11), 0.94 (3H, s, H<sub>3</sub>-19a), 0.79 (3H, s, H<sub>3</sub>-19b).  $^{13}\text{C}$  NMR (126 MHz,  $\text{DMSO}-d_6$ )  $\delta_{\text{C}}$  169.5 (C-22, C), 166.6 (C-4, C), 165.2 (C-1, C), 143.4 (C-7a, C), 143.2 (C-20, C), 136.6 (C-13, C), 132.5 (C-10a, C), 130.0 (C-15, 17, CH), 128.4 (C-8, CH), 128.1 (C-14, 18, CH), 126.5 (C-16, CH), 124.9 (C-10, CH), 124.1 (C-7, CH), 117.7 (C-9, CH), 114.1 (C-21, CH<sub>2</sub>), 78.6 (C-5a, CH), 60.5 (C-10b, CH), 58.5 (C-11b, CH), 55.5 (C-3, CH), 40.1 (C-19, C), 36.3 (C-11, CH<sub>2</sub>), 36.0 (C-12, CH<sub>2</sub>), 23.7 (C-19b, CH<sub>3</sub>), 22.9 (C-19a, CH<sub>3</sub>), 22.1 (C-23, CH<sub>3</sub>).

3-O-methylviridicatin (**7**): white powder;  $^1\text{H}$  NMR (500 MHz, Chloroform- $d$ )  $\delta_{\text{H}}$  7.53 (2H, t,  $J = 7.2$  Hz, H-2', 6'), 7.48 (1H, d,  $J = 7.2$  Hz, H-4'), 7.43 (2H, m, H-7, 8), 7.37 (2H, d,  $J = 7.2$  Hz, H-3', 5'), 7.23 (1H, d,  $J = 8.0$  Hz, H-5), 7.11 (1H, m, H-6), 3.83 (3H, s, 3-OCH<sub>3</sub>).  $^{13}\text{C}$  NMR (126 MHz,  $\text{DMSO}$ )  $\delta_{\text{C}}$  158.5 (C-2, C), 145.1 (C-3, C), 137.4 (C-4, CH), 135.8 (C-8a, C), 133.5 (C-1', C),

129.1 (C-2', CH), 129.1 (C-6', CH) 128.6 (C-7, CH), 128.4 (C-3', CH), 128.4 (C-5', CH), 128.0 (C-4', CH), 125.7 (C-5, CH), 122.0 (C-6, CH), 119.8 (C-4a, C), 115.1 (C-8, C), 59.4 (3-OCH<sub>3</sub>, CH<sub>3</sub>).

viridicatol (**8**): yellow powder; <sup>1</sup>H NMR (500 MHz, DMSO-*d*<sub>6</sub>)  $\delta_{\text{H}}$  12.19 (1H, s, 2-OH), 9.52 (1H, s, 3-OH), 9.14 (1H, s, 3'-OH), 7.35-7.27 (3H, m, H-8,5,2'), 7.13-7.04 (2H, m, H-6, 7), 6.85-6.79 (1H, m, H-4), 6.74-6.69 (2H, m, H-5',6'). <sup>13</sup>C NMR (126 MHz, DMSO)  $\delta_{\text{C}}$  158.3 (C-2, C), 157.3 (C-3', C), 142.3 (C-3, C), 134.9 (C-4, C), 133.1 (C-9, C), 129.4 (C-2', CH), 126.4 (C-5, CH), 124.4 (C-7, CH), 124.0 (C-1', C), 122.1 (C-6, CH), 120.9 (C-10, C), 120.4 (C-5', CH), 116.7 (C-6', CH), 115.2 (C-8, CH), 114.6 (C-4', CH).

arctosin (**9**): yellow oil; <sup>1</sup>H NMR (500 MHz, DMSO-*d*<sub>6</sub>)  $\delta_{\text{H}}$  7.53 (1H, ddd, *J* = 8.4, 7.5, 1.7 Hz, H-8), 7.15 (1H, dd, *J* = 8.4, 1.1 Hz, H-9), 7.09 (1H, td, *J* = 7.5, 1.1 Hz, H-7), 7.02-6.98 (1H, m, H-6), 6.96 (1H, d, *J* = 7.9 Hz, H-15), 6.69-6.65 (1H, m, H-14), 6.11 (1H, t, *J* = 2.0 Hz, H-12), 6.02-5.98 (1H, m, H-16), 4.23 (1H, s, H-10), 3.06 (3H, s, 4-CH<sub>3</sub>). <sup>13</sup>C NMR (126 MHz, DMSO-*d*<sub>6</sub>)  $\delta_{\text{C}}$  166.0 (C-2, C), 165.4 (C-5, C), 157.0 (C-13, C), 135.1 (C-9a, C), 132.4 (C-8, CH), 132.3 (C-11, C), 130.5 (C-6, CH), 128.9 (C-15, CH), 126.5 (C-5a, C), 124.2 (C-7, CH), 121.1 (C-9, CH), 116.9 (C-16, CH), 115.7 (C-14, CH), 112.8 (C-12, CH), 70.1 (C-3, C), 63.7 (C-10, C), 30.9 (4-CH<sub>3</sub>, CH<sub>3</sub>).

cyclopenin (**10**): yellow oil; [ $\alpha$ ]<sub>D</sub><sup>25</sup> -60.1 (c 0.05, MeOH), <sup>1</sup>H NMR (500 MHz, DMSO-*d*<sub>6</sub>)  $\delta_{\text{H}}$  10.84 (1H, s, -NH), 7.54 (1H, ddd, *J* = 8.4, 7.7, 1.6 Hz, H-8), 7.32-7.28 (1H, m, H-14), 7.22 (2H, t, *J* = 7.7 Hz, H-13, 15), 7.16 (1H, dd, *J* = 8.4, 1.1 Hz, H-9), 7.09 (1H, td, *J* = 7.7, 1.1 Hz, H-7), 6.92 (1H, dd, *J* = 7.7, 1.6 Hz, H-6), 6.66-6.61 (2H, m, H-12, 16), 4.36 (1H, s, H-10), 3.32 (3H, s, 4-CH<sub>3</sub>). <sup>13</sup>C NMR (126 MHz, DMSO-*d*<sub>6</sub>)  $\delta_{\text{C}}$  165.9 (C-2, C), 165.3 (C-5, C), 135.2 (C-9a, C), 132.4 (C-8, CH), 131.0 (C-11, C), 130.5 (C-9, CH), 128.7 (C-14, CH), 127.9 (C-13, CH), 127.9 (C-15, CH), 126.4 (C-5a, C), 126.1 (C-12, CH), 126.1 (C-16, CH), 124.2 (C-7, CH), 121.1 (C-6, CH), 70.1 (C-3, C), 63.7 (C-10, CH), 30.8 (4-CH<sub>3</sub>, CH<sub>3</sub>).

8-Hydroxyhelvafuranone (**12**): green oil; [ $\alpha$ ]<sub>D</sub><sup>25</sup> + 3.07 (c 0.05, MeOH), <sup>1</sup>H NMR (500 MHz, DMSO-*d*<sub>6</sub>)  $\delta_{\text{H}}$  9.33 (1H, s, 8-OH), 9.22 (1H, s, 9-OH), 9.15 (1H, s, 16-OH), 7.80 (1H, d, *J* = 8.3 Hz, 5-OH), 6.99 (2H, d, *J* = 8.1 Hz, H-14, 18), 6.93 (1H, d, *J* = 2.0 Hz, H-7), 6.80 (1H, d, *J* = 8.1 Hz, H-10), 6.76 (1H, dd, *J* = 8.2, 2.0 Hz, H-11), 6.70 (2H, d, *J* = 8.2 Hz, H-15, 17), 5.78 (1H, d, *J* = 7.7 Hz, H-5), 3.85 (1H, d, *J* = 15.1 Hz, H-12), 3.58 (1H, d, *J* = 15.1 Hz, H-12). <sup>13</sup>C NMR (126 MHz, DMSO-*d*<sub>6</sub>)  $\delta_{\text{C}}$  170.6 (C-2, C), 157.7 (C-4, C), 156.1 (C-16, C), 146.1 (C-9, C), 145.1 (C-8,

C), 129.6 (C-14, 18, CH), 127.8 (C-6, C), 126.5 (C-13, C), 120.3 (C-3, C), 120.2 (C-11, CH), 116.1 (C-7, CH), 115.6 (C-10, CH), 115.5 (C-15, 17, CH), 96.5 (C-5, CH), 31.0 (C-12, CH<sub>2</sub>).

verrucosidinol acetate (**13**): white oil;  $[\alpha]_D^{25} +9.7$  (c 0.03, MeOH), <sup>1</sup>H NMR (500 MHz, DMSO-*d*<sub>6</sub>)  $\delta_H$  5.75 (1H, d, *J* = 3.2 Hz, H-9), 5.69 (1H, s, H-7), 5.38 (1H, s, H-11), 3.96 (1H, d, *J* = 6.9, 3.2 Hz, H-15), 3.74 (3H, t, *J* = 4.2 Hz, H-24), 3.55 (1H, d, *J* = 3.2 Hz, H-13), 2.17 (3H, t, *J* = 4.2 Hz, H-25), 2.00 (3H, d, *J* = 3.2 Hz, H-17), 1.89 (3H, d, *J* = 3.4 Hz, H-16), 1.78 (3H, d, *J* = 3.0 Hz, H-20), 1.73 (3H, d, *J* = 3.0 Hz, H-19), 1.44 (3H, d, *J* = 3.2 Hz, H-18), 1.38 (3H, d, *J* = 3.2 Hz, H-22), 1.23 (3H, d, *J* = 3.1 Hz, H-21), 1.05 (3H, dd, *J* = 7.2, 3.1 Hz, H-23). <sup>13</sup>C NMR (126 MHz, DMSO)  $\delta_C$  169.2 (C-26, C), 168.4 (C-3, C), 163.6 (C-1, C), 158.4 (C-5, C), 133.8 (C-9, CH), 133.7 (C-10, C), 132.8 (C-11, CH), 130.9 (C-8, C), 111.6 (C-4, C), 109.3 (C-2, C), 81.6 (C-7, CH), 79.3 (C-12, C), 77.5 (C-6, C), 75.8 (C-15, CH), 66.9 (C-13, CH), 66.3 (C-14, C), 60.2 (C-24, C), 23.3 (C-18, CH<sub>3</sub>), 21.9 (C-21, CH<sub>3</sub>), 20.8 (C-25, CH<sub>3</sub>), 18.6 (C-23, CH<sub>3</sub>), 18.1 (C-22, CH<sub>3</sub>), 15.1 (C-19, CH<sub>3</sub>), 13.5 (C-22, CH<sub>3</sub>), 9.9 (C-16, CH<sub>3</sub>), 9.8 (C-17, CH<sub>3</sub>).

deoxyverrucosidin (**14**): yellow oil;  $[\alpha]_D^{25} -0.1$  (c 0.1, MeOH), <sup>1</sup>H NMR (500 MHz, CDCl<sub>3</sub>)  $\delta_H$  6.15 (1H, s, H-7), 5.79 (1H, s, H-9), 5.36 (1H, s, H-11), 4.10 (1H, q, *J* = 6.8 Hz, H-15), 3.81 (3H, s, H-18), 3.40 (1H, s, H-13), 2.07 (3H, s, H-17), 2.00 (3H, d, *J* = 1.5 Hz, H-21), 1.86 (3H, d, *J* = 1.5 Hz, H-22), 1.82 (3H, s, H-19), 1.66 (3H, d, *J* = 1.4 Hz, H-20), 1.15 (3H, d, *J* = 6.8 Hz, H-16). <sup>13</sup>C NMR (126 MHz, CDCl<sub>3</sub>)  $\delta_C$  168.2 (C-3, C), 166.1 (C-1, C), 156.9 (C-5, C), 137.9 (C-7, CH), 136.9 (C-9, CH), 135.1 (C-8, C), 133.6 (C-11, CH), 132.6 (C-10, C), 126.6 (C-6, C), 110.2 (C-2, C-4, 2C), 80.2 (C-12, C), 76.5 (C-15, CH), 67.6 (C-13, CH), 67.4 (C-14, C), 60.4 (C-18, CH<sub>3</sub>), 23.4 (C-23, CH<sub>3</sub>), 22.0 (C-16, CH<sub>3</sub>), 19.0 (C-21, CH<sub>3</sub>), 18.6 (C-22, CH<sub>3</sub>), 16.4 (C-20, CH<sub>3</sub>), 13.9 (C-24, CH<sub>3</sub>), 10.9 (C-19, CH<sub>3</sub>), 10.5 (C-17, CH<sub>3</sub>).

nordeoxyverrucosidin (**15**): yellow oil;  $[\alpha]_D^{25} +2.7$  (c 0.1, MeOH), <sup>1</sup>H NMR (500 MHz, CDCl<sub>3</sub>)  $\delta$  6.07 (1H, s, H-7), 5.86 (1H, s, H-9), 5.51 (1H, d, *J* = 1.5 Hz, H-11), 5.48 (1H, s, H-2), 4.14 (1H, q, *J* = 6.8 Hz, H-15), 3.83 (3H, s, 3-OCH<sub>3</sub>), 3.44 (1H, s, H-13), 2.07 (3H, d, *J* = 1.5 Hz, 6-CH<sub>3</sub>), 1.96 (9H, dd, *J* = 3.6, 2.1 Hz, 4-CH<sub>3</sub>, 8-CH<sub>3</sub>, 10-CH<sub>3</sub>), 1.48 (3H, s, 14-CH<sub>3</sub>), 1.43 (3H, s, 12-CH<sub>3</sub>), 1.21 (3H, d, *J* = 6.8 Hz, 15-CH<sub>3</sub>). <sup>13</sup>C NMR (126 MHz, CDCl<sub>3</sub>)  $\delta_C$  171.5 (C-3, C), 164.4, (C-1, C) 161.5 (C-5, C), 139.4 (C-7, CH), 136.5 (C-9, CH), 135.1 (C-8, C), 133.6 (C-11, CH), 132.1 (C-10, C), 127.5 (C-6, C), 106.8 (C-4, C), 88.3 (C-2, CH), 80.3 (C-12, C), 76.9 (C-15, CH), 67.6 (C-13, CH),

67.6 (C-14, C), 56.3 (3-OCH<sub>3</sub>, CH<sub>3</sub>), 22.0 (12-CH<sub>3</sub>, CH<sub>3</sub>), 19.0 (15-CH<sub>3</sub>, CH<sub>3</sub>), 18.7 (10-CH<sub>3</sub>, CH<sub>3</sub>), 18.6 (8-CH<sub>3</sub>, CH<sub>3</sub>), 16.7 (6-CH<sub>3</sub>, CH<sub>3</sub>), 14.0 (14-CH<sub>3</sub>, CH<sub>3</sub>), 11.2 (4-CH<sub>3</sub>, CH<sub>3</sub>).

aspterric acid methyl ester (**16**): yellow oil;  $[\alpha]_D^{25}$  -9.23 (c 0.1, MeOH) <sup>1</sup>H NMR (500 MHz, Chloroform-d)  $\delta_H$  4.17 (1H, d,  $J$  = 8.4 Hz, H-2), 3.79 (3H, s, H-16), 3.37 (1H, d,  $J$  = 8.2 Hz, H-13a), 2.46-2.40 (1H, m, H-13b), 2.37-2.32 (2H, m, H-4), 2.28 (1H, d,  $J$  = 8.3 Hz, H-6), 2.18 (1H, dd,  $J$  = 13.2, 8.4 Hz, H-1a), 2.04 (1H, d,  $J$  = 13.3 Hz, 8.4 Hz, H-1b), 2.01 (1H, d,  $J$  = 5.5 Hz, H-9a), 1.84-1.76 (1H, m, H-5a), 1.74 (3H, s, H-14), 1.70 (1H, d,  $J$  = 8.3 Hz, H-5b), 1.60 (3H, s, H-15), 1.47 (1H, d,  $J$  = 5.5 Hz, H-9b), 1.18 (1H, dt,  $J$  = 14.0, 6.6 Hz, H-8a), 1.03 (1H, d,  $J$  = 14.0, 6.6 Hz, H-8b). <sup>13</sup>C NMR (126 MHz, Chloroform-d)  $\delta_C$  176.2 (C-12, C), 135.5 (C-10, C), 124.6 (C-11, C), 84.9 (C-2, CH), 78.6 (C-3, C), 76.2 (C-13, CH<sub>2</sub>), 55.7 (C-6, CH), 53.1 (C-7, C), 52.6 (C-16, CH<sub>3</sub>), 36.3 (C-1, CH<sub>2</sub>), 34.4 (C-4, CH<sub>2</sub>), 33.4 (C-8, CH<sub>2</sub>), 32.4 (C-9, CH<sub>2</sub>), 23.7 (C-5, CH<sub>2</sub>), 23.5 (C-15, CH<sub>3</sub>), 21.0 (C-14, CH<sub>3</sub>).