

Supporting Information

Synthesis of Piperine Analogs Containing Isoxazoline/Pyrazoline Scaffold and Their Pesticidal Bioactivities

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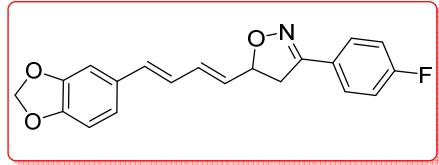
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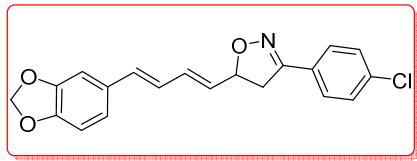
Content

1 Characterization.....	2-14
2 Bioassay pictures.....	15-16
3 ¹HNMR and ¹³C NMR Spectra.....	17-76

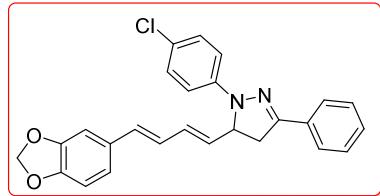
1 Characterization



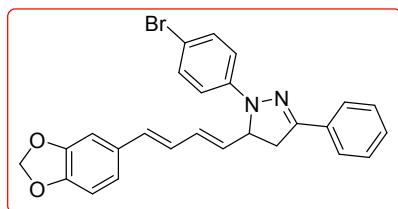
Data for VIb: Yield = 5%; Light yellow solid, m.p. 130–132 °C. IR cm⁻¹ (KBr): 3010, 2900, 1605, 1504, 1490, 1443, 1383, 1252, 1040, 982, 841; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.72-7.74 (m, 2H), 7.28-7.31 (m, 2H), 7.15 (s, 1H), 6.86-6.92 (m, 2H), 6.78-6.83 (m, 1H, H-3'), 6.61 (d, *J* = 15.5 Hz, 1H, H-4'), 6.45-6.50 (m, 1H, H-2'), 6.01 (s, 2H, OCH₂O), 5.91 (dd, *J* = 7.5 Hz, *J* = 15.0 Hz, 1H, H-1'), 5.20-5.25 (m, 1H, H-5), 3.62 (dd, *J* = 10.5 Hz, 16.5 Hz, 1H, H-4), 3.25 (dd, *J* = 9.0 Hz, 17.0 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 164.4 (*J* = 246.2 Hz), 156.4, 148.3, 147.6, 133.89, 133.86, 131.7, 131.1, 129.4 (*J* = 7.5 Hz), 126.6, 126.5, 122.1, 116.4 (*J* = 21.2 Hz), 108.8, 105.8, 101.6, 82.1, 40.3. HRMS (ESI): Calcd for C₂₀H₁₇O₃NF ([M+H]⁺), 338.1187; found, 338.1186.



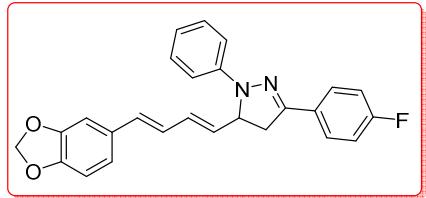
Data for VIc: Yield = 13%; White solid, m.p. 132–134 °C. IR cm⁻¹ (KBr): 3010, 2920, 1504, 1492, 1448, 1254, 1096, 1045, 984, 822; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.70 (d, *J* = 9.0 Hz, 2H), 7.54 (d, *J* = 9.0 Hz, 2H), 7.16 (s, 1H), 6.87-6.92 (m, 2H), 6.84 (dd, *J* = 10.5 Hz, 15.5 Hz, 1H, H-3'), 6.62 (d, *J* = 15.5 Hz, 1H, H-4'), 6.50 (dd, *J* = 10.5 Hz, 15.0 Hz, 1H, H-2'), 6.02 (s, 2H, OCH₂O), 5.91 (dd, *J* = 7.5 Hz, 15.0 Hz, 1H, H-1'), 5.22-5.27 (m, 1H, H-5), 3.62 (dd, *J* = 10.5 Hz, 17.0 Hz, 1H, H-4), 3.25 (dd, *J* = 9.0 Hz, 17.0 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 156.5, 148.3, 147.6, 135.0, 133.9, 131.7, 131.0, 129.3, 128.5, 128.8, 126.6, 122.1, 108.8, 105.8, 101.6, 82.3, 40.3. HRMS (ESI): Calcd for C₂₀H₁₇O₃NCl ([M+H]⁺), 354.0891; found, 354.0891.



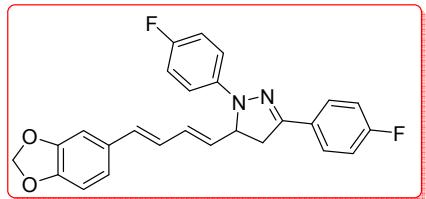
Data for VIIc: Yield = 55%; Yellow solid, m.p. 100–102 °C. IR cm⁻¹ (KBr): 3016, 2880, 1594, 1501, 1488, 1253, 1040, 981, 815, 684; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.75 (d, *J* = 7.5 Hz, 2H), 7.45 (t, *J* = 7.5 Hz, 2H), 7.37-7.40 (m, 1H), 7.28 (d, *J* = 9.0 Hz, 2H), 7.15 (d, *J* = 9.0 Hz, 2H), 7.10 (s, 1H), 6.84-6.88 (m, 2H), 6.77 (dd, *J* = 10.5 Hz, 15.5 Hz, 1H, H-3’), 6.53 (d, *J* = 15.5 Hz, 1H, H-4’), 6.44 (dd, *J* = 10.5 Hz, 15.0 Hz, 1H, H-2’), 6.00 (s, 2H, OCH₂O), 5.81 (dd, *J* = 7.5 Hz, 15.0 Hz, 1H, H-1’), 5.02-5.07 (m, 1H, H-5), 3.68 (dd, *J* = 11.5 Hz, 17.0 Hz, 1H, H-4), 3.17 (dd, *J* = 5.5 Hz, 17.0 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 149.3, 148.3, 147.5, 143.9, 133.1, 132.66, 132.61, 132.0, 131.7, 129.1, 126.8, 126.3, 126.2, 122.7, 121.9, 115.2, 108.8, 105.7, 101.6, 101.5, 62.0, 40.4. HRMS (ESI): Calcd for C₂₆H₂₂O₂N₂Cl ([M+H]⁺), 429.1364; found, 429.1366.



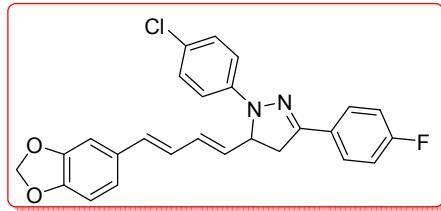
Data for VIIId: Yield = 77%; Yellow solid, m.p. 102–104 °C. IR cm⁻¹ (KBr): 3029, 2887, 1588, 1501, 1488, 1250, 1038, 985, 816, 691; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.74-7.75 (m, 2H), 7.45 (t, *J* = 7.5 Hz, 2H), 7.37-7.40 (m, 3H), 7.08-7.10 (m, 3H), 6.84-6.88 (m, 2H), 6.77 (dd, *J* = 10.5 Hz, 16.0 Hz, 1H, H-3’), 6.53 (d, *J* = 16.0 Hz, 1H, H-4’), 6.44 (dd, *J* = 10.5 Hz, 15.5 Hz, 1H, H-2’), 5.99 (s, 2H, OCH₂O), 5.81 (dd, *J* = 7.5 Hz, 15.0 Hz, 1H, H-1’), 5.02-5.07 (m, 1H, H-5), 3.68 (dd, *J* = 11.5 Hz, 17.0 Hz, 1H, H-4), 3.17 (dd, *J* = 5.5 Hz, 17.0 Hz, 1H, H-4). HRMS (ESI): Calcd for C₂₆H₂₂O₂N₂Br ([M+H]⁺), 473.0859; found, 473.0859.



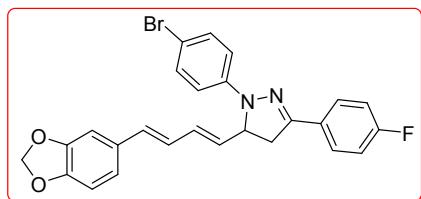
Data for VIIe: Yield = 30%; Yellow solid, m.p. 108–110 °C. IR cm⁻¹ (KBr): 3015, 2890, 1596, 1497, 1446, 1251, 1038, 985, 833, 691; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.77-7.79 (m, 2H), 7.21-7.29 (m, 4H), 7.15 (d, *J* = 8.0 Hz, 2H), 7.10 (s, 1H), 6.84-6.88 (m, 2H), 6.72-6.78 (m, 2H), 6.53 (d, *J* = 15.5 Hz, 1H, H-4'), 6.46 (dd, *J* = 10.5 Hz, 15.0 Hz, 1H, H-2'), 5.99 (s, 2H, OCH₂O), 5.83 (dd, *J* = 7.5 Hz, 15.0 Hz, 1H, H-1'), 5.00-5.05 (m, 1H, H-5), 3.66 (dd, *J* = 11.5 Hz, 17.5 Hz, 1H, H-4), 3.15 (dd, *J* = 6.0 Hz, 17.0 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 163.7 (*J* = 245.0 Hz), 148.2, 147.7, 147.4, 145.2, 132.9, 132.6, 132.4, 131.8, 129.5, 129.3, 128.2 (*J* = 8.7 Hz), 126.9, 121.9, 119.2, 116.2 (*J* = 21.2 Hz), 113.9, 108.8, 105.6, 101.5, 62.3, 40.3. HRMS (ESI): Calcd for C₂₆H₂₂O₂N₂F ([M+H]⁺), 413.1660; found, 413.1659.



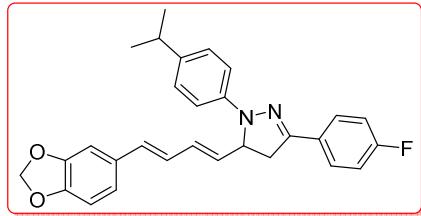
Data for VIIIf: Yield = 61%; Light yellow solid, m.p. 143–145 °C. IR cm⁻¹ (KBr): 3009, 2920, 1504, 1446, 1255, 1043, 985, 821; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.76-7.79 (m, 2H), 7.29 (t, *J* = 8.5 Hz, 2H), 7.13-7.15 (m, 2H), 7.06-7.10 (m, 3H), 6.84-6.88 (m, 2H), 6.78 (dd, *J* = 10.5 Hz, 15.5 Hz, 1H, H-3'), 6.53 (d, *J* = 15.5 Hz, 1H, H-4'), 6.47 (dd, *J* = 10.5 Hz, 15.0 Hz, 1H, H-2'), 6.00 (s, 2H, OCH₂O), 5.82 (dd, *J* = 7.5 Hz, 15.0 Hz, 1H, H-1'), 4.96-5.01 (m, 1H, H-5), 3.66 (dd, *J* = 11.5 Hz, 17.0 Hz, 1H, H-4), 3.15 (dd, *J* = 6.5 Hz, 17.0 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 163.8 (*J* = 246.2 Hz), 157.4 (*J* = 232.5 Hz), 148.3, 148.0, 147.4, 142.1, 133.0, 132.7, 132.4, 131.7, 129.4, 128.3 (*J* = 7.5 Hz), 126.8, 121.9, 116.2 (*J* = 21.2 Hz), 115.9 (*J* = 22.5 Hz), 115.2 (*J* = 7.5 Hz), 108.8, 105.7, 101.5, 63.0, 40.6. HRMS (ESI): Calcd for C₂₆H₂₁O₂N₂F₂ ([M+H]⁺), 431.1566; found, 431.1564.



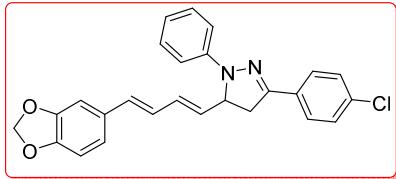
Data for VIIg: Yield = 65%; Light yellow solid, m.p. 152–154 °C. IR cm⁻¹ (KBr): 3015, 2962, 1601, 1514, 1492, 1447, 1255, 1124, 983, 829; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.77–7.80 (m, 2H), 7.25–7.29 (m, 4H), 7.14 (d, *J* = 8.5 Hz, 2H), 7.10 (s, 1H), 6.84–6.88 (m, 2H), 6.77 (dd, *J* = 10.5 Hz, 15.5 Hz, 1H, H-3’), 6.53 (d, *J* = 15.5 Hz, 1H, H-4’), 6.45 (dd, *J* = 11.0 Hz, 15.0 Hz, 1H, H-2’), 6.00 (s, 2H, OCH₂O), 5.80 (dd, *J* = 7.5 Hz, 15.0 Hz, 1H, H-1’), 5.02–5.07 (m, 1H, H-5), 3.68 (dd, *J* = 11.5 Hz, 17.0 Hz, 1H, H-4), 3.17 (dd, *J* = 5.5 Hz, 17.5 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 163.9 (*J* = 245.0 Hz), 148.5, 148.3, 147.5, 144.0, 133.1, 132.6, 132.1, 131.8, 129.3 (*J* = 2.5 Hz), 129.1, 128.4, 126.8, 122.8, 122.0, 116.2 (*J* = 21.2 Hz), 115.3, 108.8, 105.7, 101.6, 62.3, 40.3. HRMS (ESI): Calcd for C₂₆H₂₁O₂N₂ClF ([M+H]⁺), 447.1270; found, 447.1269.



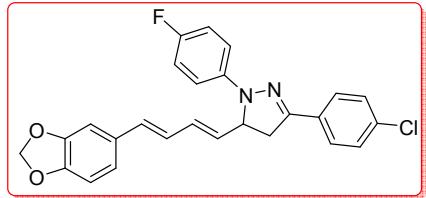
Data for VIIh: Yield = 50%; Light yellow solid, m.p. 125–127 °C. IR cm⁻¹ (KBr): 3002, 2920, 1715, 1490, 1445, 1363, 1222, 1039, 987, 834; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.77–7.80 (m, 2H), 7.39 (d, *J* = 8.5 Hz, 2H), 7.26–7.29 (m, 2H), 7.07–7.10 (m, 3H), 6.84–6.88 (m, 2H), 6.77 (dd, *J* = 10.5 Hz, 15.5 Hz, 1H, H-3’), 6.53 (d, *J* = 15.5 Hz, 1H, H-4’), 6.44 (dd, *J* = 10.5 Hz, 15.0 Hz, 1H, H-2’), 6.00 (s, 2H, OCH₂O), 5.80 (dd, *J* = 7.5 Hz, 15.0 Hz, 1H, H-1’), 5.02–5.07 (m, 1H, H-5), 3.68 (dd, *J* = 11.5 Hz, 17.0 Hz, 1H, H-4), 3.17 (dd, *J* = 5.5 Hz, 17.5 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 163.9 (*J* = 245.0 Hz), 148.6, 148.2, 147.5, 144.2, 133.1, 132.8, 132.6, 131.9, 131.7, 131.7, 129.2, 128.4 (*J* = 7.5 Hz), 126.8, 121.9, 116.2 (*J* = 21.2 Hz), 115.7, 110.3, 108.8, 105.7, 101.5, 62.0, 40.3. HRMS (ESI): Calcd for C₂₆H₂₁O₂N₂BrF ([M+H]⁺), 491.0765; found, 491.0765.



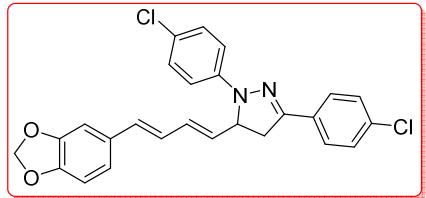
Data for VIIi: Yield = 34%; Yellow solid, m.p. 160–162 °C. IR cm⁻¹ (KBr): 3018, 2957, 2922, 1610, 1508, 1488, 1446, 1251, 1038, 985, 833; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.74-7.77 (m, 2H), 7.28 (t, *J* = 9.0 Hz, 2H), 7.06-7.11 (m, 5H), 6.84-6.88 (m, 2H), 6.78 (dd, *J* = 10.5 Hz, 15.5 Hz, 1H, H-3'), 6.54 (d, *J* = 16.0 Hz, 1H, H-4'), 6.47 (dd, *J* = 10.5 Hz, 15.0 Hz, 1H, H-2'), 6.00 (s, 2H, OCH₂O), 5.84 (dd, *J* = 7.5 Hz, 15.0 Hz, 1H, H-1'), 4.94-4.99 (m, 1H, H-5), 3.64 (dd, *J* = 11.5 Hz, 17.0 Hz, 1H, H-4), 3.12 (dd, *J* = 6.5 Hz, 17.0 Hz, 1H, H-4), 2.76-2.81 (m, 1H, CH₃CHCH₃), 1.16 (d, *J* = 7.0 Hz, 6H, CH₃CHCH₃). HRMS (ESI): Calcd for C₂₉H₂₆O₂N₂F ([M-H]⁺), 453.1973; found, 453.1973.



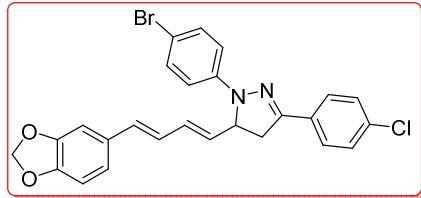
Data for VIIj: Yield = 42%; Light yellow solid, m.p. 135–137 °C. IR cm⁻¹ (KBr): 3023, 2917, 1597, 1500, 1490, 1445, 1251, 1039, 985, 745, 691; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.75 (d, *J* = 8.5 Hz, 2H), 7.50 (d, *J* = 9.0 Hz, 2H), 7.22-7.25 (m, 2H), 7.16 (d, *J* = 8.0 Hz, 2H), 7.10 (d, *J* = 0.5 Hz, 1H), 6.84-6.88 (m, 2H), 6.72-6.79 (m, 2H), 6.53 (d, *J* = 16.0 Hz, 1H, H-4'), 6.46 (dd, *J* = 10.5 Hz, 15.5 Hz, 1H, H-2'), 5.99 (s, 2H, OCH₂O), 5.83 (dd, *J* = 7.5 Hz, 15.0 Hz, 1H, H-1'), 5.03-5.08 (m, 1H, H-5), 3.66 (dd, *J* = 11.5 Hz, 17.0 Hz, 1H, H-4), 3.14 (dd, *J* = 6.0 Hz, 17.0 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 148.3, 147.5, 147.4, 144.9, 133.5, 133.0, 132.5, 131.85, 131.81, 129.4, 129.2, 127.8, 127.7, 126.9, 121.9, 119.4, 113.9, 108.8, 105.7, 101.5, 62.3, 40.4. HRMS (ESI): Calcd for C₂₆H₂₂O₂N₂Cl ([M+H]⁺), 429.1364; found, 429.1363.



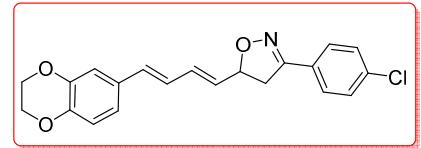
Data for VIIIk: Yield = 47%; Light yellow solid, m.p. 130–132 °C. IR cm⁻¹ (KBr): 3015, 2919, 2850, 1505, 1499, 1446, 1255, 1040, 985, 822; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.74 (d, *J* = 8.5 Hz, 2H), 7.49 (d, *J* = 8.0 Hz, 2H), 7.07-7.16 (m, 5H), 6.84-6.88 (m, 2H), 6.77 (dd, *J* = 10.5 Hz, 15.5 Hz, 1H, H-3'), 6.53 (d, *J* = 15.5 Hz, 1H, H-4'), 6.47 (dd, *J* = 10.5 Hz, 15.0 Hz, 1H, H-2'), 6.00 (s, 2H, OCH₂O), 5.82 (dd, *J* = 7.5 Hz, 15.0 Hz, 1H, H-1'), 4.99-5.04 (m, 1H, H-5), 3.65 (dd, *J* = 11.5 Hz, 17.0 Hz, 1H, H-4), 3.14 (dd, *J* = 6.5 Hz, 17.0 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 157.5 (*J* = 233.7 Hz), 148.3, 147.7, 147.5, 141.9, 133.6, 133.1, 132.8, 132.3, 131.79, 131.78, 129.2, 127.8, 126.8, 122.0, 116.0 (*J* = 22.5 Hz), 115.3 (*J* = 6.2 Hz), 108.8, 105.6, 101.6, 62.9, 40.3. HRMS (ESI): Calcd for C₂₆H₂₁O₂N₂ClF ([M+H]⁺), 447.1270; found, 447.1270.



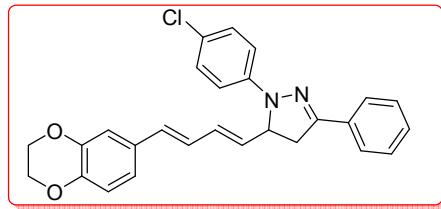
Data for VIIIL: Yield = 68%; Light yellow solid, m.p. 117–119 °C. IR cm⁻¹ (KBr): 3019, 2892, 1598, 1489, 1447, 1254, 1040, 984, 821; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.76 (d, *J* = 8.0 Hz, 2H), 7.50 (d, *J* = 8.5 Hz, 2H), 7.28 (d, *J* = 8.5 Hz, 2H), 7.15 (d, *J* = 8.5 Hz, 2H), 7.10 (s, 1H), 6.84-6.88 (m, 2H), 6.77 (dd, *J* = 10.5 Hz, 15.5 Hz, 1H, H-3'), 6.53 (d, *J* = 15.5 Hz, 1H, H-4'), 6.45 (dd, *J* = 11.0 Hz, 14.5 Hz, 1H, H-2'), 6.00 (s, 2H, OCH₂O), 5.80 (dd, *J* = 7.5 Hz, 15.0 Hz, 1H, H-1'), 5.05-5.10 (m, 1H, H-5), 3.67 (dd, *J* = 11.5 Hz, 17.0 Hz, 1H, H-4), 3.16 (dd, *J* = 5.0 Hz, 17.5 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 148.3, 148.2, 147.5, 143.7, 133.7, 133.1, 132.7, 131.9, 131.7, 131.5, 129.18, 129.16, 127.8, 126.7, 122.9, 121.9, 115.3, 108.8, 105.7, 101.5, 62.6, 40.3. HRMS (ESI): Calcd for C₂₆H₂₁O₂N₂Cl₂ ([M+H]⁺), 463.0975; found, 463.0975.



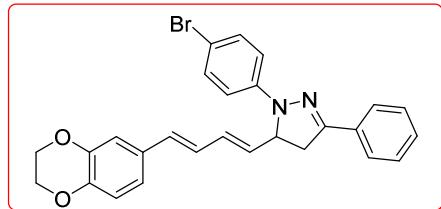
Data for VIIIm: Yield = 76%: Yellow solid, m.p. 114–116 °C. IR cm⁻¹ (KBr): 3024, 2897, 1587, 1500, 1488, 1446, 1251, 1040, 987, 817; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.76 (d, *J* = 8.5 Hz, 2H), 7.50 (d, *J* = 8.5 Hz, 2H), 7.39 (d, *J* = 9.0 Hz, 2H), 7.08-7.10 (m, 3H), 6.84-6.88 (m, 2H), 6.76 (dd, *J* = 10.5 Hz, 15.5 Hz, 1H, H-3'), 6.53 (d, *J* = 15.5 Hz, 1H, H-4'), 6.44 (dd, *J* = 10.5 Hz, 15.5 Hz, 1H, H-2'), 6.00 (s, 2H, OCH₂O), 5.80 (dd, *J* = 7.5 Hz, 15.0 Hz, 1H, H-1'), 5.05-5.10 (m, 1H, H-5), 3.67 (dd, *J* = 11.5 Hz, 17.0 Hz, 1H, H-4), 3.17 (dd, *J* = 5.5 Hz, 17.5 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 148.4, 148.3, 147.5, 144.0, 133.8, 133.2, 132.7, 132.0, 131.8, 131.7, 131.6, 129.2, 127.9, 126.8, 122.0, 115.8, 110.6, 108.9, 105.7, 101.6, 62.1, 40.4. HRMS (ESI): Calcd for C₂₆H₂₁O₂N₂BrCl ([M+H]⁺), 507.0469; found, 507.0469.



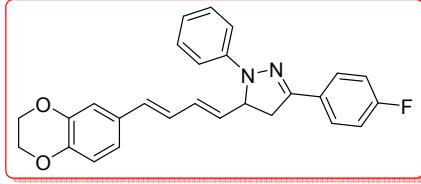
Data for VIIIc: Yield = 36%: Light yellow solid, m.p. 207–209 °C. IR cm⁻¹ (KBr): 3008, 2976, 2910, 2870, 1583, 1500, 1299, 1126, 1070, 984, 861, 668; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.70 (d, *J* = 8.5 Hz, 2H), 7.54 (d, *J* = 8.5 Hz, 2H), 6.99 (d, *J* = 2.0 Hz, 1H), 6.96 (dd, *J* = 2.0 Hz, 8.5 Hz, 1H), 6.75-6.82 (m, 2H), 6.57 (d, *J* = 10.5 Hz, 1H, H-4'), 6.50 (dd, *J* = 10.5 Hz, 15.0 Hz, 1H, H-2'), 5.91 (dd, *J* = 7.5 Hz, 15.0 Hz, 1H, H-1'), 5.26 (dd, *J* = 8.5 Hz, 18.5 Hz, 1H, H-5), 4.23 (s, 4H, OCH₂CH₂O), 3.62 (dd, *J* = 10.5 Hz, 17.0 Hz, 1H, H-4), 3.25 (dd, *J* = 8.5 Hz, 16.5 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 155.9, 143.39, 143.34, 134.4, 133.3, 133.1, 130.3, 130.1, 128.7, 128.28, 128.21, 126.0, 119.7, 117.1, 114.5, 81.7, 64.0, 63.9, 39.7. HRMS (ESI): Calcd for C₂₁H₁₉ClNO₃ ([M+H]⁺), 368.1048; found, 368.1046.



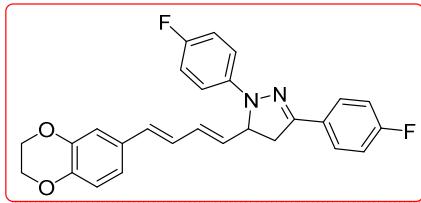
Data for IXc: Yield = 42%; Yellow solid, m.p. 94–96 °C. IR cm⁻¹ (KBr): 3020, 2922, 2872, 1594, 1504, 1489, 1391, 1288, 1068, 987, 819, 758, 691; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.75 (d, *J* = 7.0 Hz, 2H), 7.45 (t, *J* = 7.0 Hz, 2H), 7.37-7.39 (m, 1H), 7.27 (d, *J* = 8.5 Hz, 2H), 7.15 (d, *J* = 8.0 Hz, 2H), 6.94 (s, 1H), 6.92 (d, *J* = 8.5 Hz, 1H), 6.79 (d, *J* = 8.0 Hz, 1H), 6.74 (dd, *J* = 10.5 Hz, 15.5 Hz, 1H), 6.39-6.48 (m, 2H), 5.82 (dd, *J* = 7.5 Hz, 15.0 Hz, 1H, H-1'), 5.02-5.07 (m, 1H, H-5), 4.21 (s, 4H, OCH₂CH₂O), 3.68 (dd, *J* = 11.5 Hz, 17.0 Hz, 1H, H-4), 3.17 (dd, *J* = 5.0 Hz, 17.0 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 149.3, 143.9, 143.7, 132.8, 132.6, 132.0, 130.8, 129.3, 129.1, 126.8, 126.2, 122.7, 120.1, 117.6, 115.2, 115.0, 64.6, 64.4, 62.0, 40.3. HRMS (ESI): Calcd for C₂₇H₂₄O₂N₂Cl ([M+H]⁺), 443.1521; found, 443.1520.



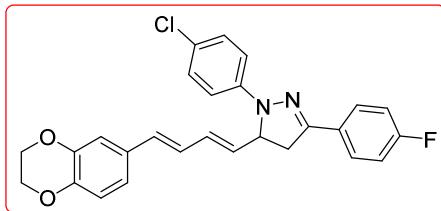
Data for IXd: Yield = 34%; Yellow solid, m.p. 106–108 °C. IR cm⁻¹ (KBr): 3004, 2922, 2871, 1587, 1505, 1487, 1391, 1288, 1068, 987, 816, 758, 691; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.75 (d, *J* = 7.0 Hz, 2H), 7.37-7.43 (m, 5H), 7.10 (d, *J* = 8.5 Hz, 2H), 6.94 (s, 1H), 6.91 (d, *J* = 8.0 Hz, 1H), 6.79 (d, *J* = 8.0 Hz, 1H), 6.74 (dd, *J* = 11.0 Hz, 15.0 Hz, 1H), 6.38-6.48 (m, 2H), 5.81 (dd, *J* = 7.5 Hz, 15.0 Hz, 1H, H-1'), 5.02-5.07 (m, 1H, H-5), 4.21 (s, 4H, OCH₂CH₂O), 3.68 (dd, *J* = 12.0 Hz, 17.0 Hz, 1H, H-4), 3.17 (dd, *J* = 4.5 Hz, 17.0 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 149.4, 144.2, 143.9, 143.7, 132.8, 132.6, 132.5, 131.9, 130.8, 129.4, 129.1, 126.8, 126.2, 120.1, 117.6, 115.7, 115.0, 110.3, 64.6, 64.4, 61.9, 40.3. HRMS (ESI): Calcd for C₂₇H₂₄O₂N₂Br ([M+H]⁺), 487.1016; found, 487.1015.



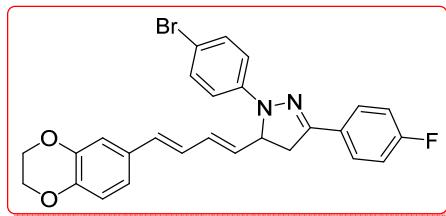
Data for IXe: Yield = 38%; Yellow solid, m.p. 88–90 °C. IR cm⁻¹ (KBr): 3008, 2923, 2878, 1596, 1497, 1288, 1067, 987, 748; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.78 (brs, 2H), 7.22-7.27 (m, 4H), 7.14-7.15 (m, 2H), 6.90-6.94 (m, 2H), 6.69-6.77 (m, 3H), 6.41-6.48 (m, 2H), 5.80-5.82 (m, 1H, H-1'), 5.02 (brs, 1H, H-5), 4.21 (s, 4H, OCH₂CH₂O), 3.60-3.66 (m, 1H, H-4), 3.14 (d, *J* = 16.0 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 163.8 (*J* = 245.0 Hz), 147.7, 145.2, 143.9, 143.7, 132.6, 132.5, 132.4, 130.8, 129.5, 129.3, 128.2 (*J* = 7.5 Hz), 126.9, 120.1, 119.2, 117.6, 116.2 (*J* = 21.2 Hz), 115.0, 113.9, 64.6, 64.5, 62.3, 40.3. HRMS (ESI): Calcd for C₂₇H₂₄FN₂O₂ ([M+H]⁺), 427.1816; found, 427.1811.



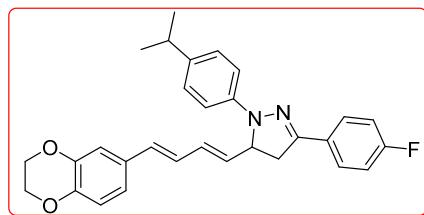
Data for IXf: Yield = 75%; Yellow solid, m.p. 93–95 °C. IR cm⁻¹ (KBr): 3002, 2918, 2873, 1504, 1288, 1220, 1068, 988, 833, 668; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.78 (brs, 2H), 7.27 (brs, 2H), 7.08-7.13 (m, 4H), 6.92-6.95 (m, 2H), 6.70-6.79 (m, 2H), 6.41-6.49 (m, 2H), 5.81 (brs, 1H, H-1'), 4.98 (brs, 1H, H-5), 4.22 (s, 4H, OCH₂CH₂O), 3.61-3.66 (m, 1H, H-4), 3.14 (d, *J* = 15.0 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 163.8 (*J* = 245.0 Hz), 157.5 (*J* = 233.7 Hz), 147.9, 143.9, 143.7, 142.2, 132.78, 132.70, 132.3, 130.8, 129.4, 128.3 (*J* = 21.2 Hz), 126.8, 120.1, 117.6, 116.1 (*J* = 21.2 Hz), 115.9 (*J* = 21.2 Hz), 115.2 (*J* = 21.2 Hz), 115.0, 64.6, 64.5, 63.0, 40.3. HRMS (ESI): Calcd for C₂₇H₂₃F₂N₂O₂ ([M+H]⁺), 445.1722; found, 445.1731.



Data for IXg: Yield = 41%; Yellow solid, m.p. 90–92 °C. IR cm⁻¹ (KBr): 3001, 2924, 2878, 1597, 1491, 1288, 1068, 987, 833, 668; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.79 (brs, 2H), 7.26-7.27 (m, 4H), 7.14 (d, *J* = 7.0 Hz, 2H), 6.90-6.94 (m, 2H), 6.69-6.79 (m, 2H), 6.39-6.48 (m, 2H), 5.77-5.81 (m, 1H, H-1'), 5.04 (brs, 1H, H-5), 4.21 (s, 4H, OCH₂CH₂O), 3.62-3.67 (m, 1H, H-4), 3.17 (d, *J* = 18.0 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 163.9 (*J* = 245.0 Hz), 148.5, 143.96, 143.91, 143.7, 132.8, 132.6, 131.9, 130.8, 129.2, 129.1, 128.4 (*J* = 8.7 Hz), 126.8, 122.7, 120.1, 117.6, 116.2 (*J* = 22.5 Hz), 115.2, 115.0, 64.6, 64.4, 62.1, 40.3. HRMS (ESI): Calcd for C₂₇H₂₃ClFN₂O₂ ([M+H]⁺), 461.1426; found, 461.1417.

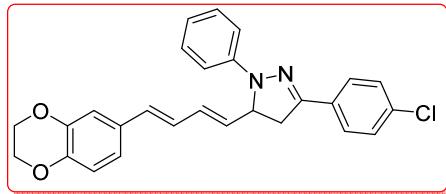


Data for IXh: Yield = 59%; Pale white solid, m.p. 99–101 °C. IR cm⁻¹ (KBr): 3005, 2923, 2882, 1590, 1508, 1489, 1382, 1289, 1219, 1069, 986, 835; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.79 (brs, 2H), 7.38 (d, *J* = 7.5 Hz, 2H), 7.26-7.29 (m, 2H), 7.09 (d, *J* = 7.5 Hz, 2H), 6.90-6.94 (m, 2H), 6.68-6.79 (m, 2H), 6.38-6.48 (m, 2H), 5.76-5.80 (m, 1H, H-1'), 5.04 (brs, 1H, H-5), 4.21 (s, 4H, OCH₂CH₂O), 3.62-3.67 (m, 1H, H-4), 3.16 (d, *J* = 13.5 Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ: 163.9 (*J* = 241.2 Hz), 148.6, 144.2, 143.9, 143.7, 132.8, 132.6, 131.9, 130.8, 129.3, 129.2, 128.4 (*J* = 5.0 Hz), 126.8, 120.1, 117.6, 116.2 (*J* = 21.2 Hz), 115.7, 115.0, 110.3, 64.6, 64.4, 62.0, 40.3. HRMS (ESI): Calcd for C₂₇H₂₃BrFN₂O₂ ([M+H]⁺), 505.0921; found, 505.0903.

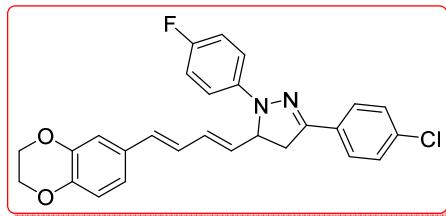


Data for IXi: Yield = 27%; Yellow solid, m.p. 84–86 °C. IR cm⁻¹ (KBr): 3002, 2955, 2923, 2869, 1609, 1507, 1288, 1067, 987, 833, 668; ¹H NMR (500 MHz, DMSO-*d*₆) δ: 7.76 (brs, 2H), 7.26 (brs, 2H), 7.08 (brs, 4H), 6.91-6.94 (m, 2H), 6.69-6.78 (m, 2H), 6.46-6.49 (m, 2H), 5.83 (brs, 1H, H-1'), 4.96 (brs, 1H, H-5), 4.21 (s, 4H, OCH₂CH₂O),

3.58-3.63 (m, 1H, H-4), 3.10 (d, J = 12.0 Hz, 1H, H-4), 2.77 (brs, 1H, CH_3CHCH_3), 1.15 (s, 6H, CH_3CHCH_3). ^{13}C NMR (125 MHz, DMSO- d_6) δ : 163.7 (J = 245.0 Hz), 147.1, 143.9, 143.7, 143.4, 139.3, 132.8, 132.6, 132.4, 130.8, 129.6, 128.1 (J = 8.7 Hz), 127.0, 126.9, 120.1, 117.6, 116.2 (J = 21.2 Hz), 115.0, 114.0, 64.6, 64.5, 62.7, 40.3, 33.0, 24.6, 24.5. HRMS (ESI): Calcd for $\text{C}_{30}\text{H}_{30}\text{FN}_2\text{O}_2$ ([M+H] $^+$), 469.2285; found, 469.2280.

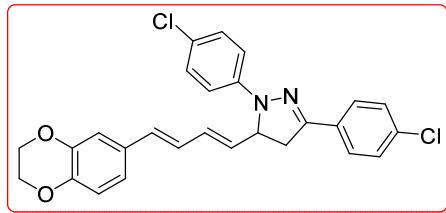


Data for IXj: Yield = 85%: Yellow solid, m.p. 145–147 °C. IR cm⁻¹ (KBr): 3014, 2972, 2932, 2885, 1599, 1502, 1288, 1070, 986, 777, 668; ^1H NMR (500 MHz, DMSO- d_6) δ : 7.74 (brs, 2H), 7.49 (brs, 2H), 7.15-7.22 (m, 4H), 6.91-6.94 (m, 2H), 6.68-6.78 (m, 3H), 6.45-6.48 (m, 2H), 5.81 (brs, 1H, H-1'), 5.05 (brs, 1H, H-5), 4.21 (s, 4H, $\text{OCH}_2\text{CH}_2\text{O}$), 3.62 (brs, 1H, H-4), 3.12 (d, J = 10.5 Hz, 1H, H-4). ^{13}C NMR (125 MHz, DMSO- d_6) δ : 147.3, 144.8, 143.8, 143.6, 133.4, 132.6, 132.3, 131.7, 130.7, 129.3, 129.1, 127.7, 126.8, 122.9, 120.1, 119.3, 117.6, 114.9, 113.8, 64.5, 64.4, 62.2, 40.2. HRMS (ESI): HRMS (ESI): Calcd for $\text{C}_{27}\text{H}_{24}\text{ClN}_2\text{O}_2$ ([M+H] $^+$), 443.1520; found, 443.1520.

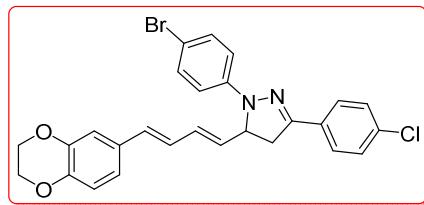


Data for IXk: Yield = 80%: Yellow solid, m.p. 91–93 °C. IR cm⁻¹ (KBr): 3017, 2980, 2923, 1579, 1507, 1288, 1216, 1067, 987, 824; ^1H NMR (500 MHz, DMSO- d_6) δ : 7.74 (brs, 2H), 7.49 (brs, 2H), 7.09-7.14 (m, 4H), 6.91-6.95 (m, 2H), 6.69-6.79 (m, 2H), 6.46-6.48 (m, 2H), 5.80 (brs, 1H, H-1'), 5.01 (brs, 1H, H-5), 4.22 (s, 4H, $\text{OCH}_2\text{CH}_2\text{O}$), 3.62 (brs, 1H, H-4), 3.13 (d, J = 12.5 Hz, 1H, H-4). ^{13}C NMR (125 MHz, DMSO- d_6) δ : 157.5 (J = 232.5 Hz), 147.7, 143.9, 143.7, 141.9, 133.5, 132.8, 132.7, 132.2, 131.7, 130.8, 129.1, 127.7, 126.8, 120.1, 117.6, 115.9 (J = 22.5 Hz),

115.3 ($J = 7.5$ Hz), 115.0, 64.6, 64.5, 63.0, 40.3. HRMS (ESI): Calcd for $C_{27}H_{23}ClFN_2O_2$ ([M+H]⁺), 461.1426; found, 461.1428.



Data for IXl: Yield = 69%; Yellow solid, m.p. 145–147 °C. IR cm⁻¹ (KBr): 3008, 2927, 2875, 1597, 1507, 1488, 1384, 1290, 1124, 1068, 985, 820; ¹H NMR (500 MHz, DMSO-*d*₆) δ : 7.78 (d, $J = 5.5$ Hz, 2H), 7.52 (d, $J = 6.0$ Hz, 2H), 7.30 (d, $J = 6.5$ Hz, 2H), 7.17 (d, $J = 6.5$ Hz, 2H), 6.92-6.97 (m, 2H), 6.71-6.81 (m, 2H), 6.41-6.51 (m, 2H), 5.79-5.83 (m, 1H, H-1'), 5.09 (brs, 1H, H-5), 4.24 (s, 4H, OCH₂CH₂O), 3.64-3.68 (m, 1H, H-4), 3.18 (d, $J = 15.0$ Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ : 148.3, 143.9, 143.79, 143.75, 133.7, 132.9, 132.7, 131.86, 131.85, 131.5, 130.8, 129.1, 127.9, 126.7, 122.9, 120.2, 117.7, 115.3, 115.0, 64.6, 64.4, 62.2, 40.3. HRMS (ESI): Calcd for $C_{27}H_{23}Cl_2N_2O_2$ ([M+H]⁺), 477.1131; found, 477.1095.



Data for IXm: Yield = 65%; Light yellow solid, m.p. 125–127 °C. IR cm⁻¹ (KBr): 3010, 2922, 2882, 1582, 1507, 1487, 1290, 1124, 1069, 986, 818; ¹H NMR (500 MHz, DMSO-*d*₆) δ : 7.75 (d, $J = 5.5$ Hz, 2H), 7.50 (d, $J = 5.0$ Hz, 2H), 7.39 (d, $J = 6.0$ Hz, 2H), 7.10 (d, $J = 6.5$ Hz, 2H), 6.90-6.94 (m, 2H), 6.68-6.79 (m, 2H), 6.38-6.48 (m, 2H), 5.80 (dd, $J = 6.5$ Hz, 13.0 Hz, 1H, H-1'), 5.07 (brs, 1H, H-5), 4.21 (s, 4H, OCH₂CH₂O), 3.61-3.67 (m, 1H, H-4), 3.15 (d, $J = 14.0$ Hz, 1H, H-4). ¹³C NMR (125 MHz, DMSO-*d*₆) δ : 148.3, 144.0, 143.9, 143.7, 133.7, 132.9, 132.6, 132.0, 131.7, 131.5, 130.8, 129.2, 127.9, 126.7, 120.2, 117.6, 115.8, 115.0, 110.5, 64.6, 64.4, 62.0, 40.3. HRMS (ESI): Calcd for $C_{27}H_{23}BrClN_2O_2$ ([M+H]⁺), 521.0625; found, 521.0600.

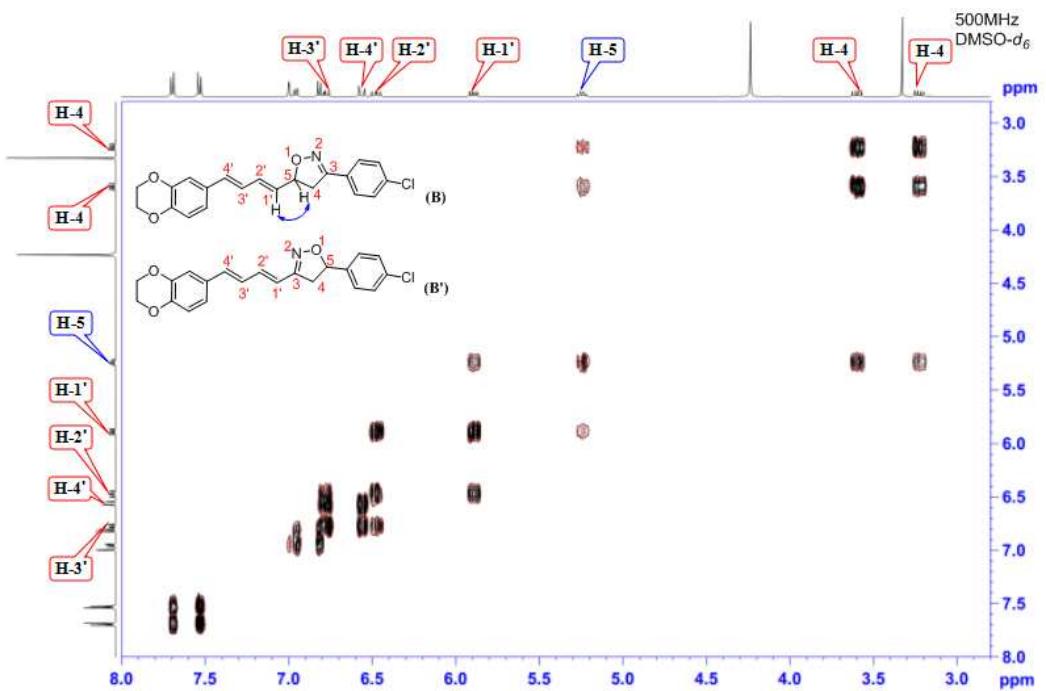


Figure S1. ^1H - ^1H COSY spectrum of compound **VIIc** (structure **B** is the right isomer).

2 Bioassay pictures



Figure S2. The representative abnormal larvae pictures of compounds **VIIb** (YRG-129), **VIa** (YRG-130), **VIII** (YRG-133), and **VIIg** (YRG-139) against *M. separata* during the larval period (CK: blank control group).

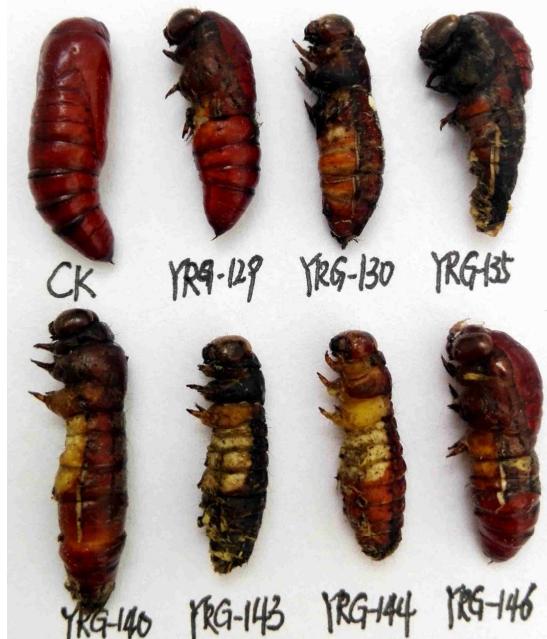


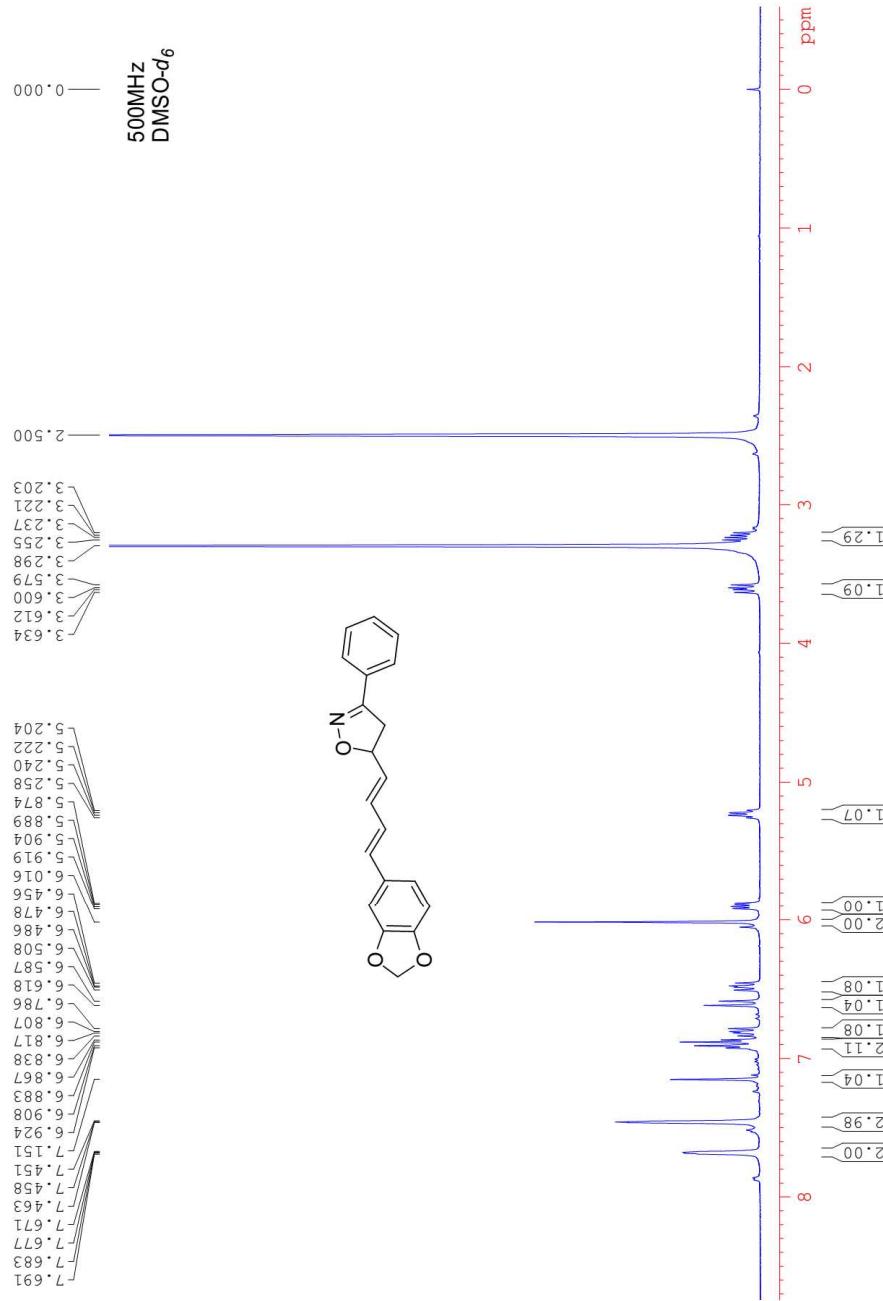
Figure S3. The representative malformed pupae pictures of compounds **VIIb** (YRG-129), **VIa** (YRG-130), **VIc** (YRG-135), **VIIh** (YRG-140), **VIIIa** (YRG-143), **VIIIb** (YRG-144), and **VIIIId** (YRG-146) against *M. separata* during the pupation period (CK: blank control group).



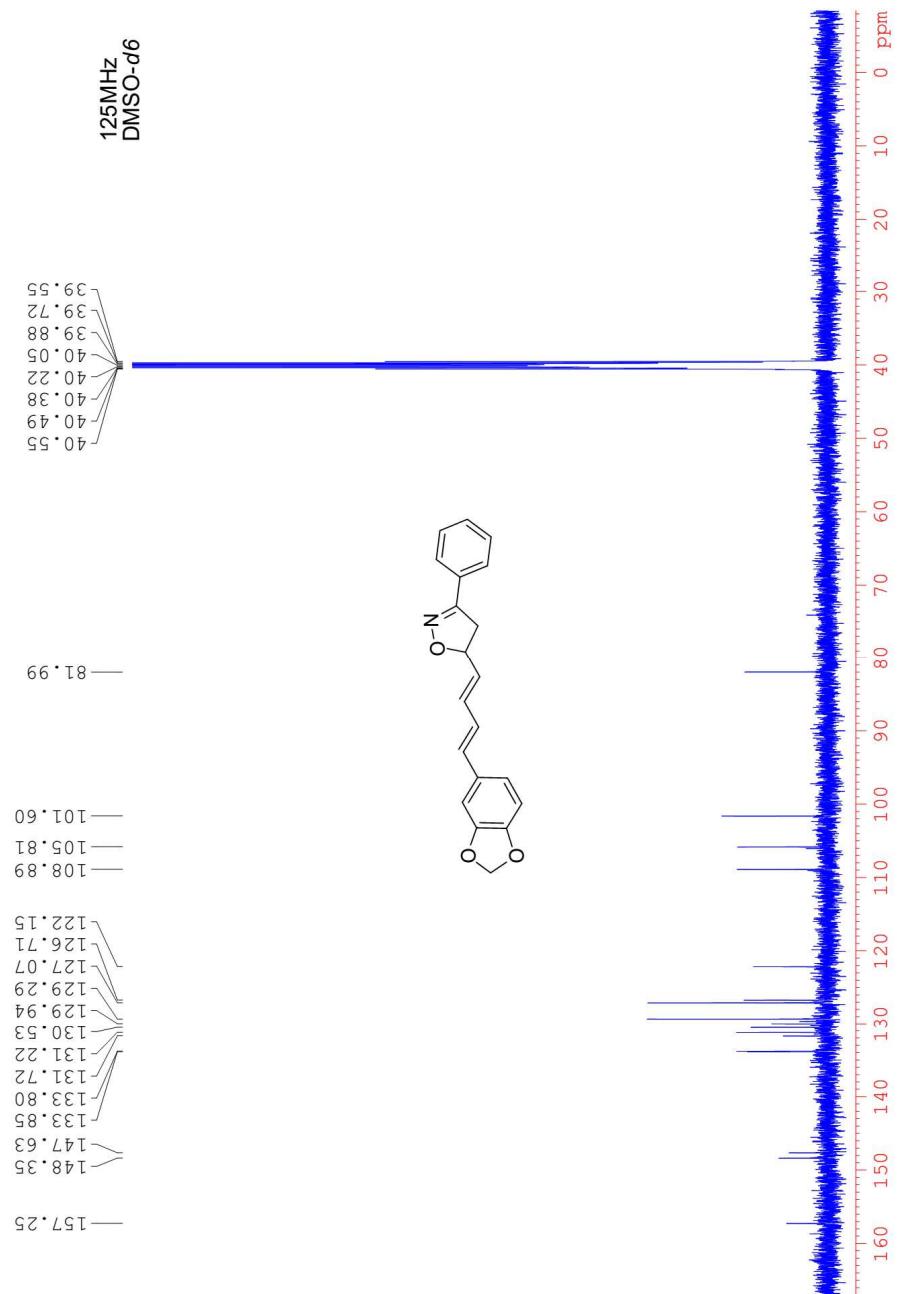
Figure S4. The representative malformed moth pictures of compounds **VIIb** (YRG-129), **VIc** (YRG-135), and **VIIh** (YRG-140) against *M. separata* during the adult period (CK: blank control group).

3 ^1H NMR and ^{13}C NMR Spectra

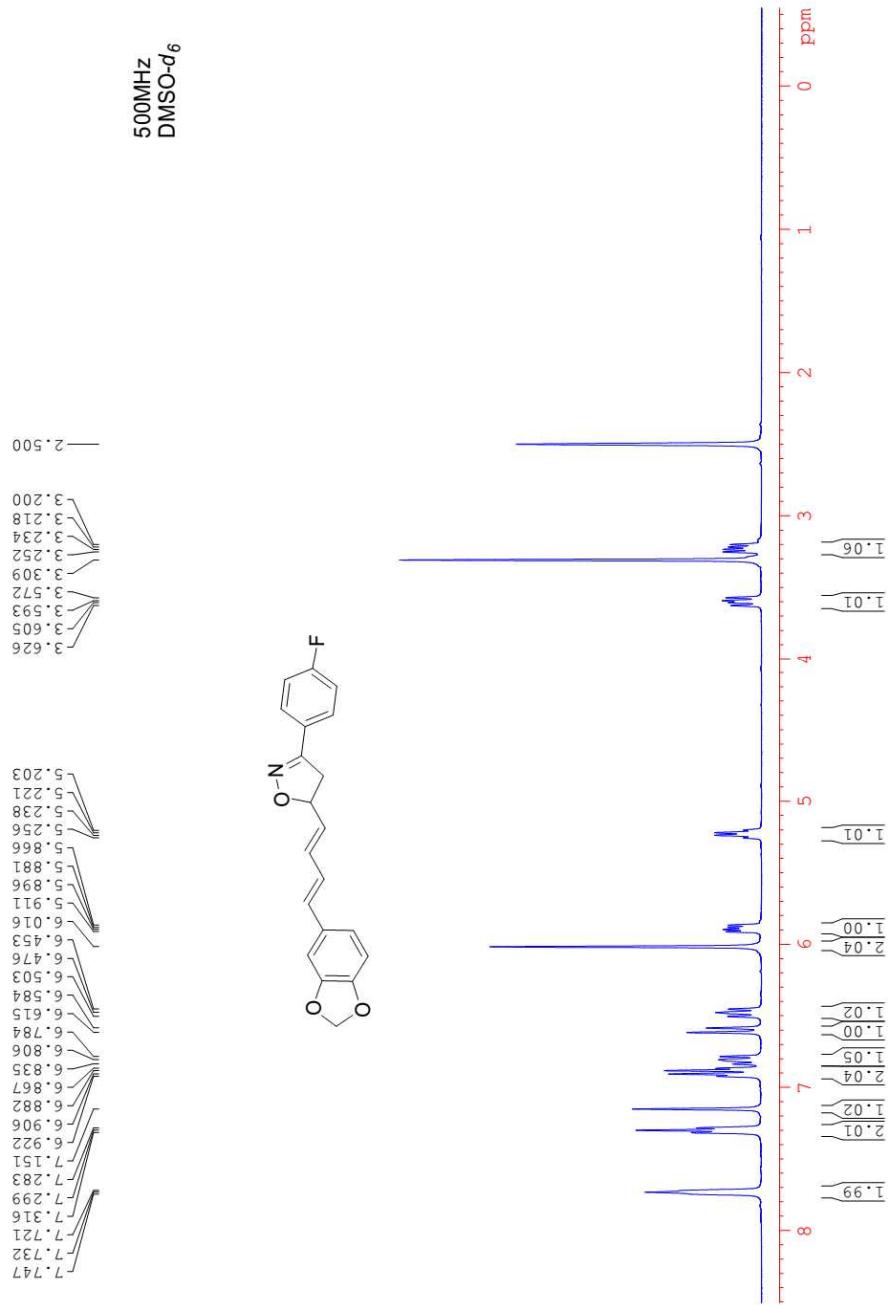
compound VIa



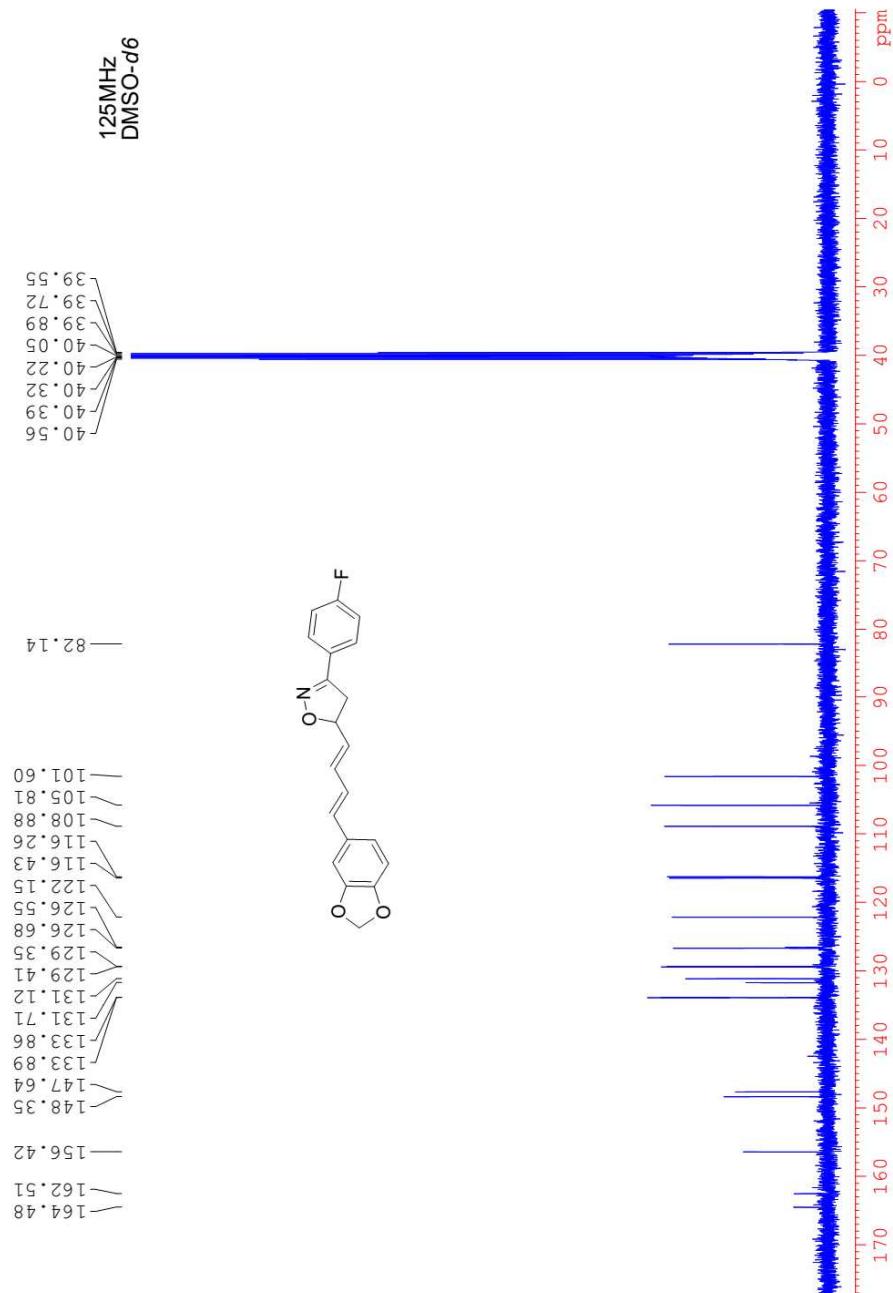
compound VIa



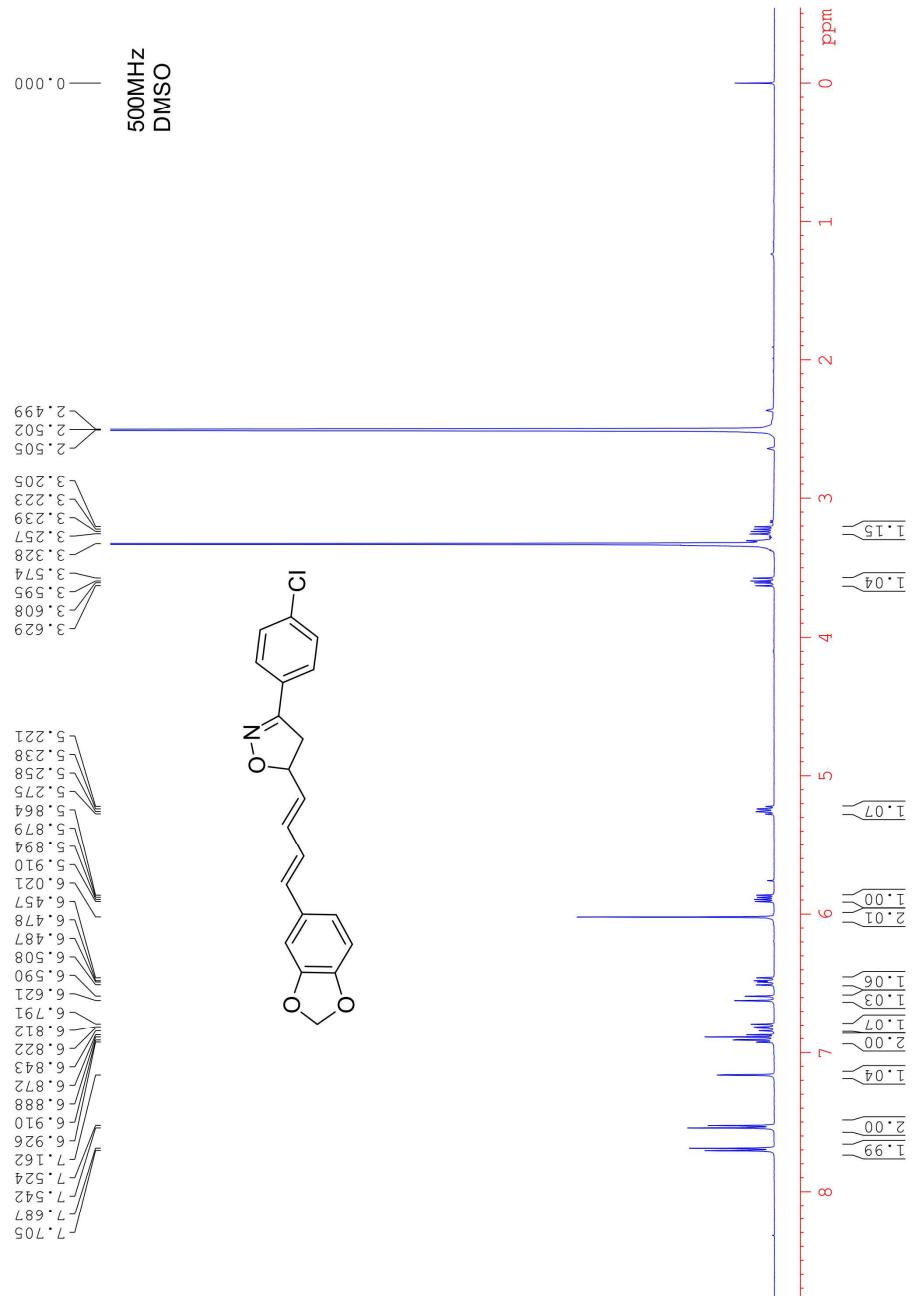
compound VIb



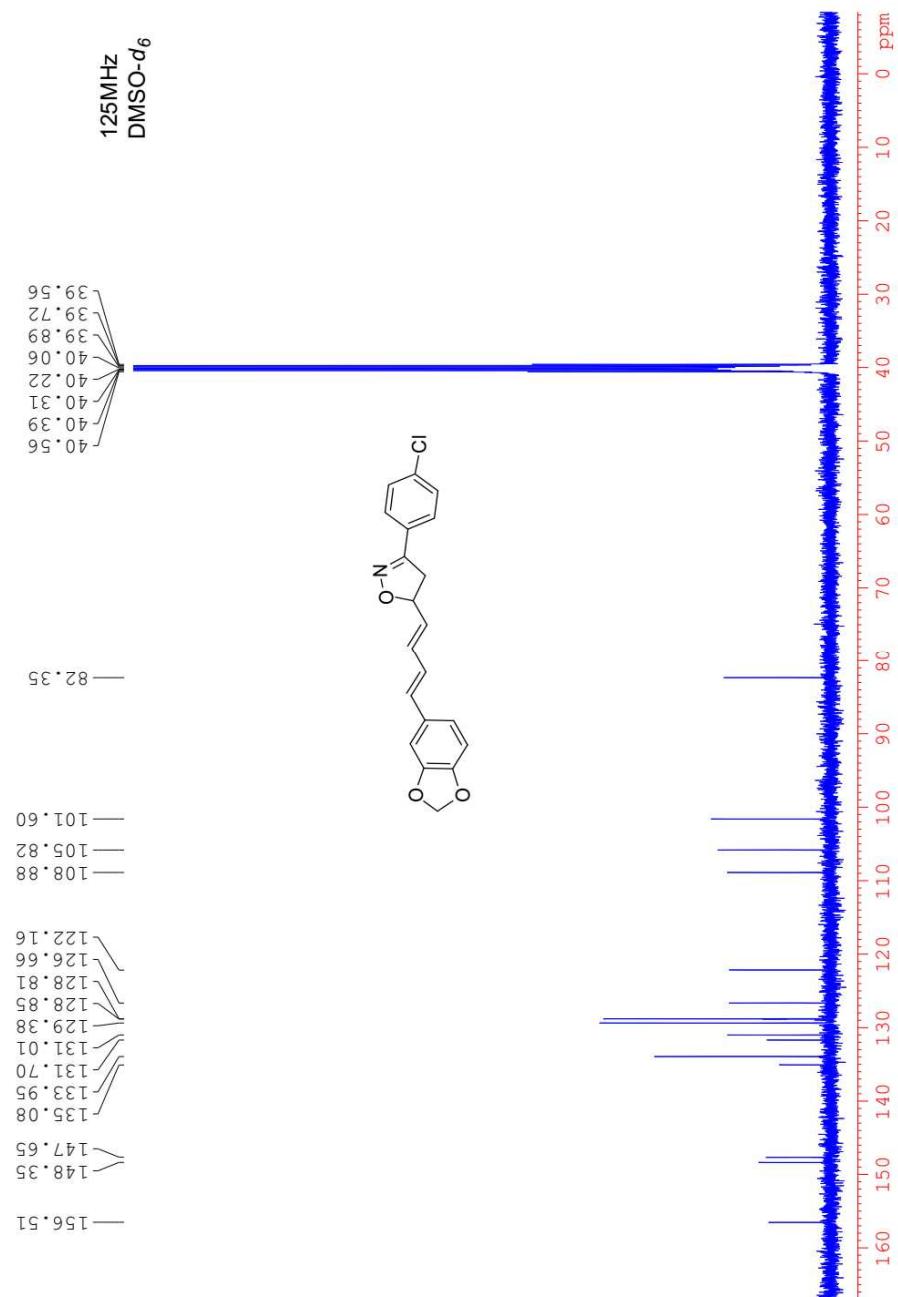
compound VIb



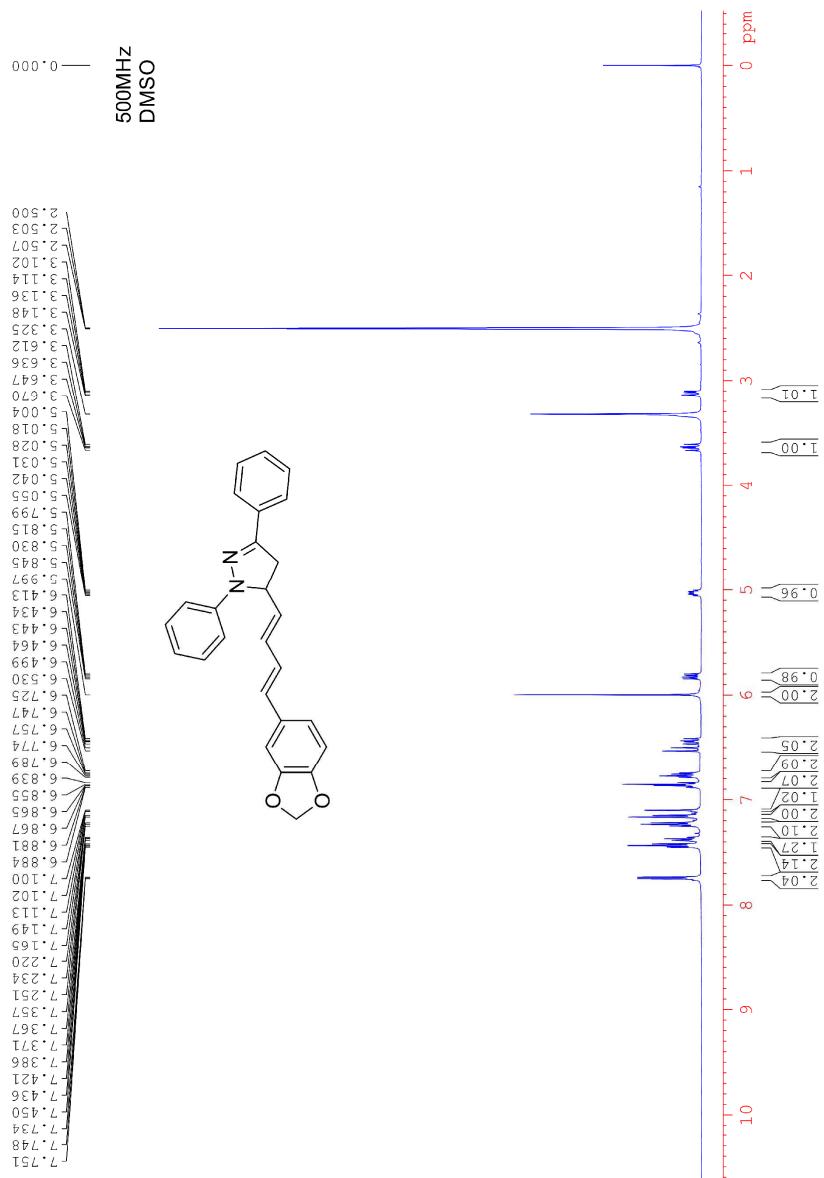
compound VIc



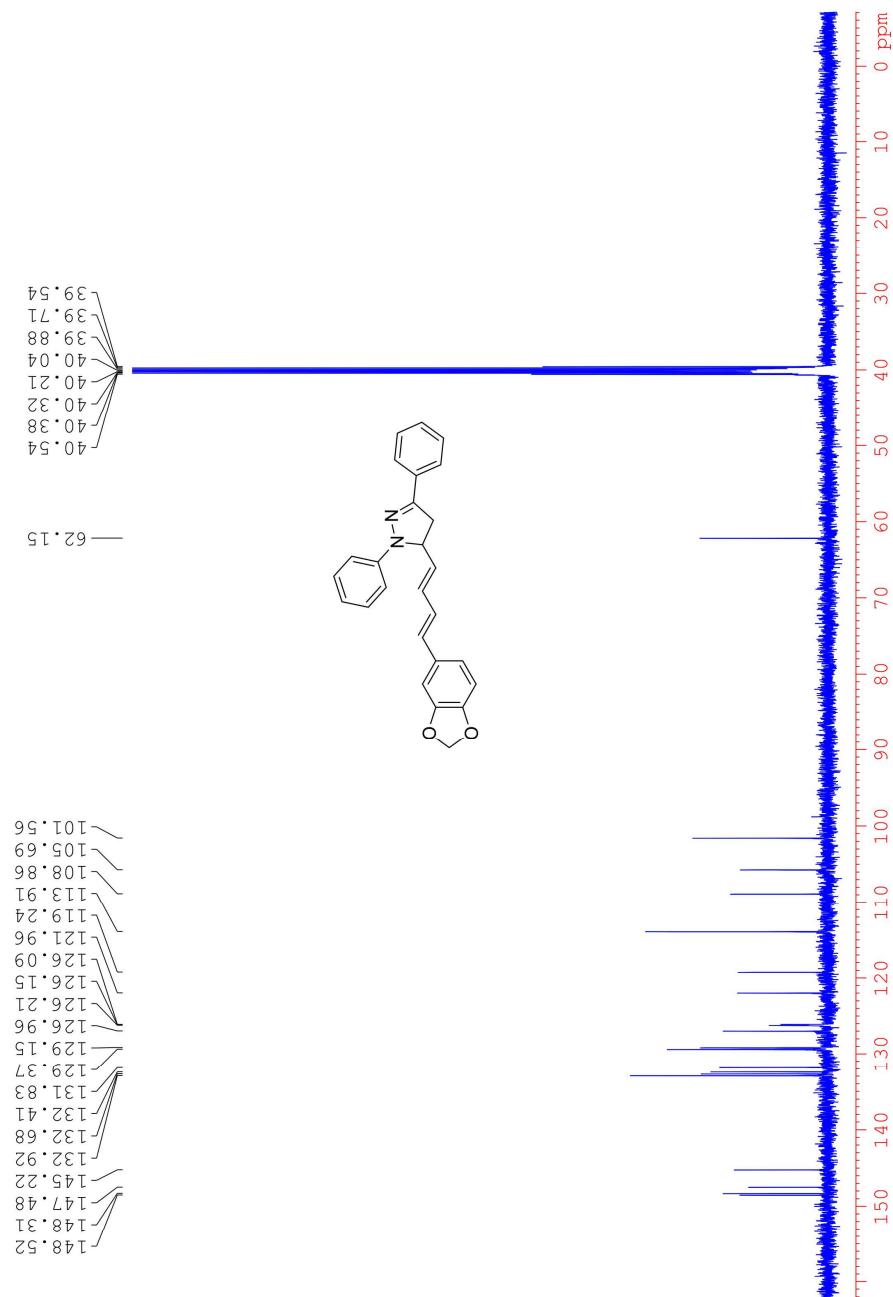
compound VIc



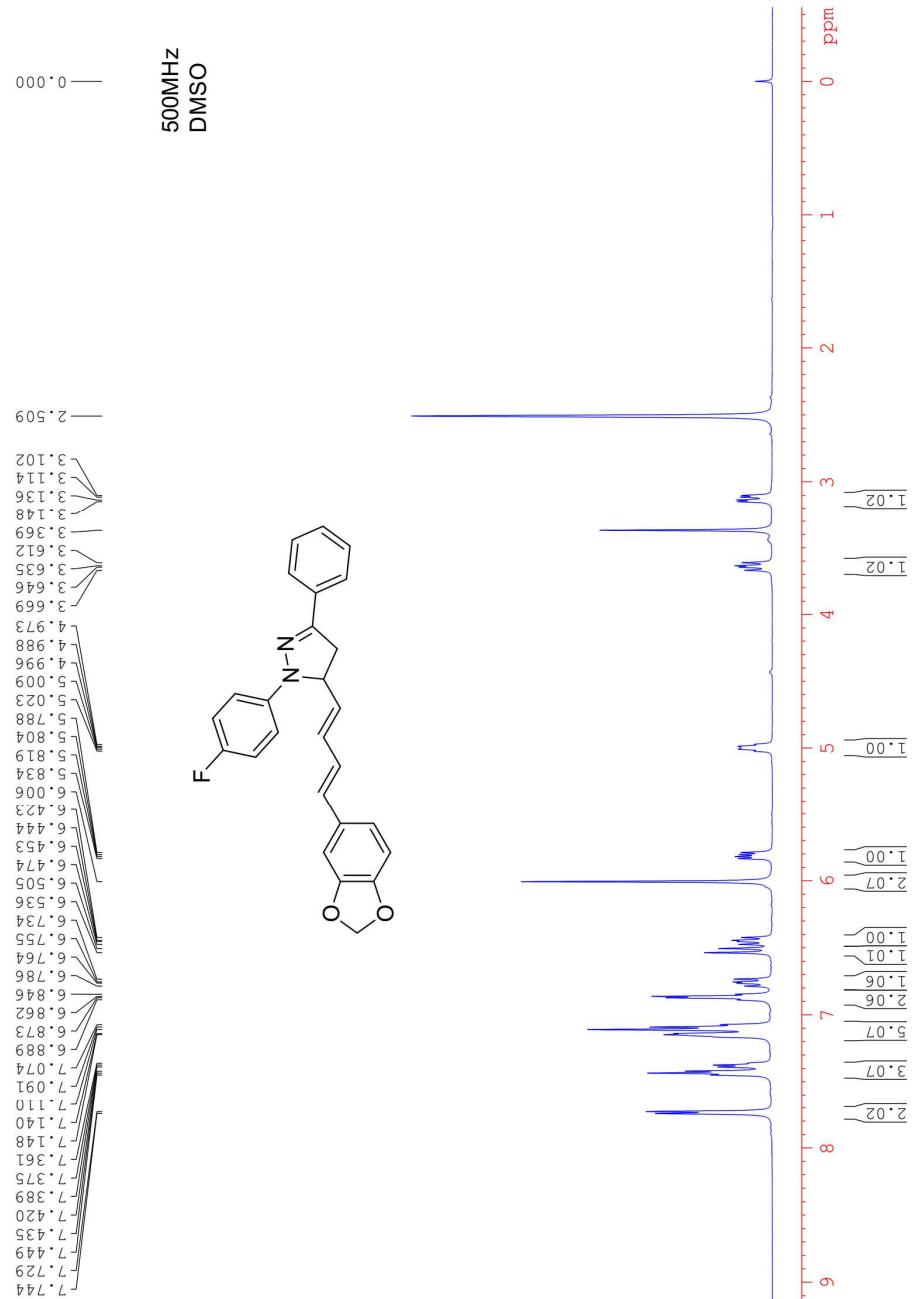
compound VIIa



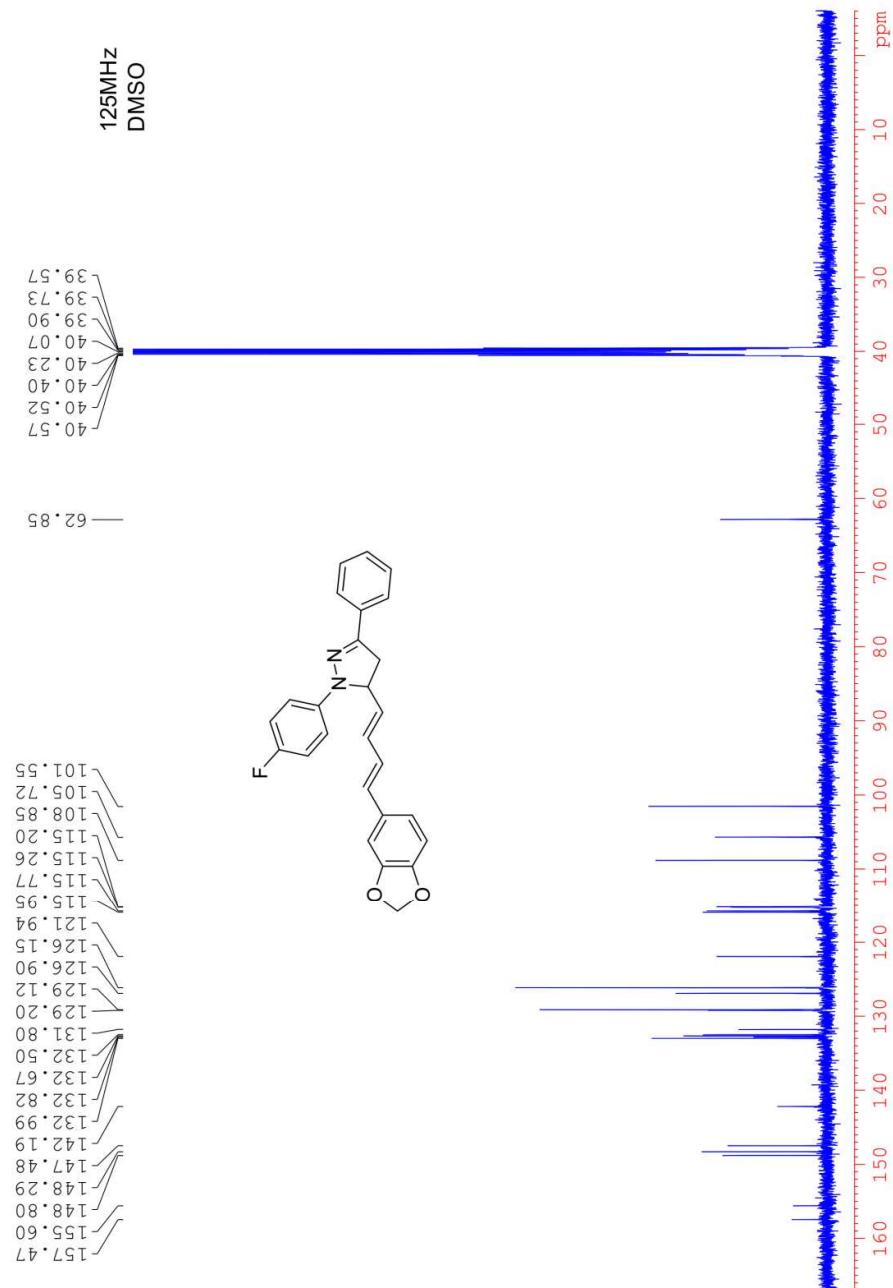
compound VIIa



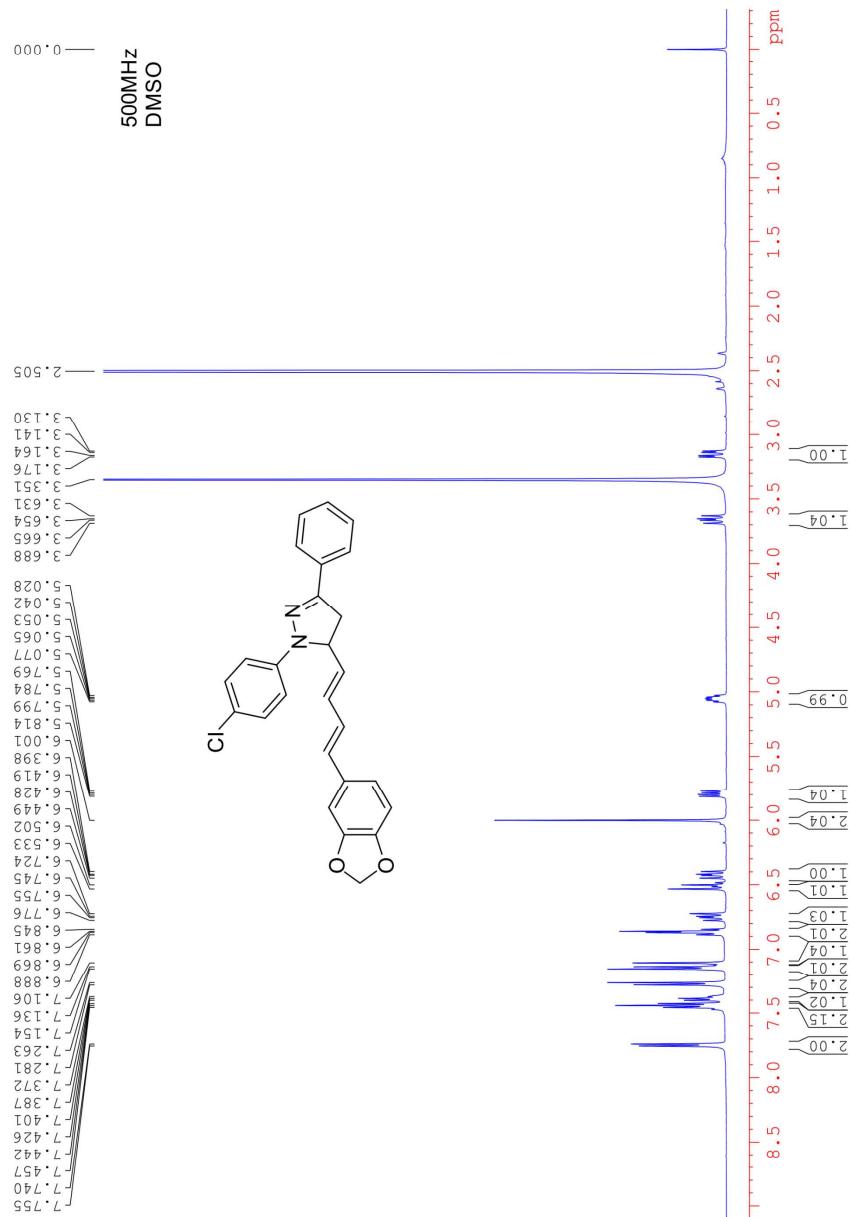
compound VIIb



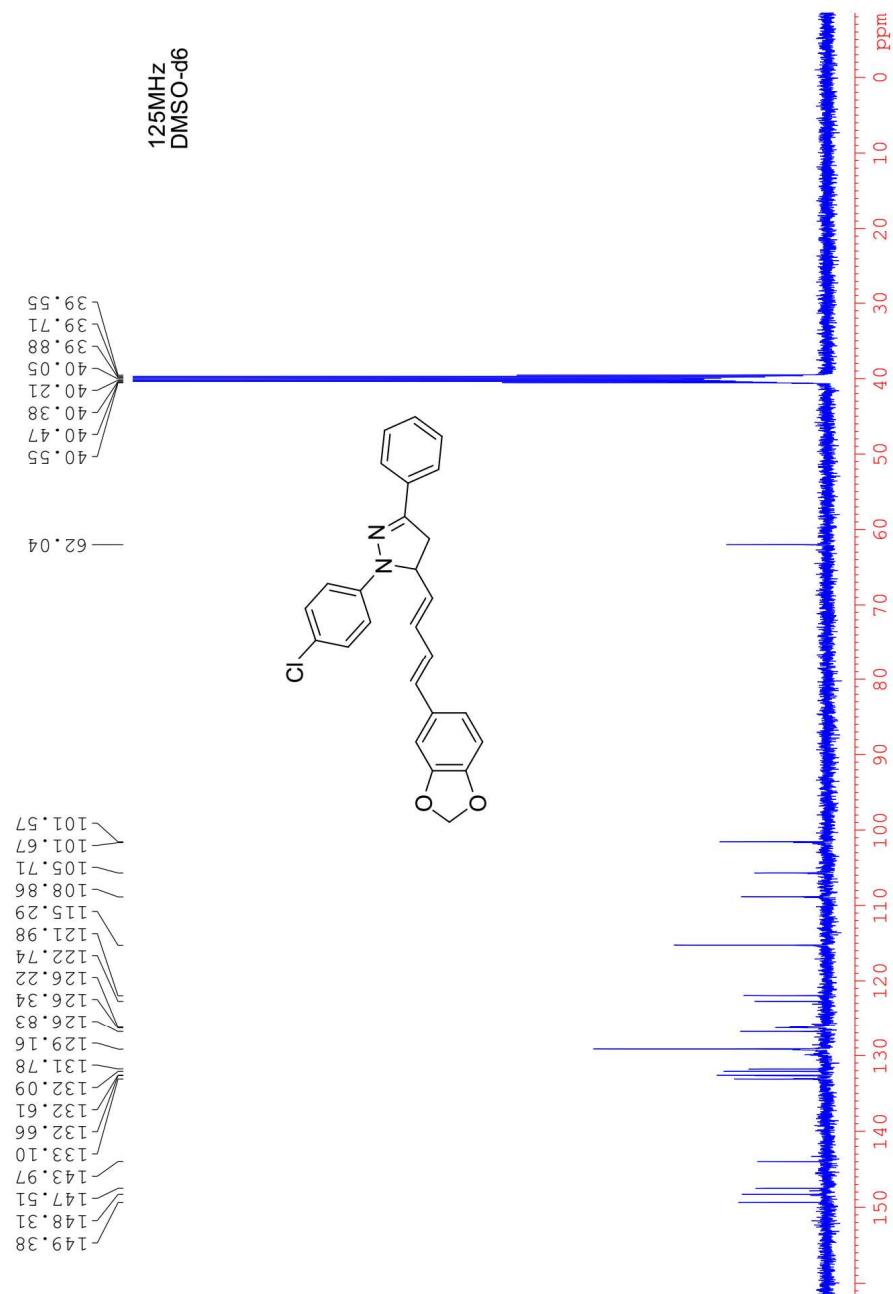
compound VIIb



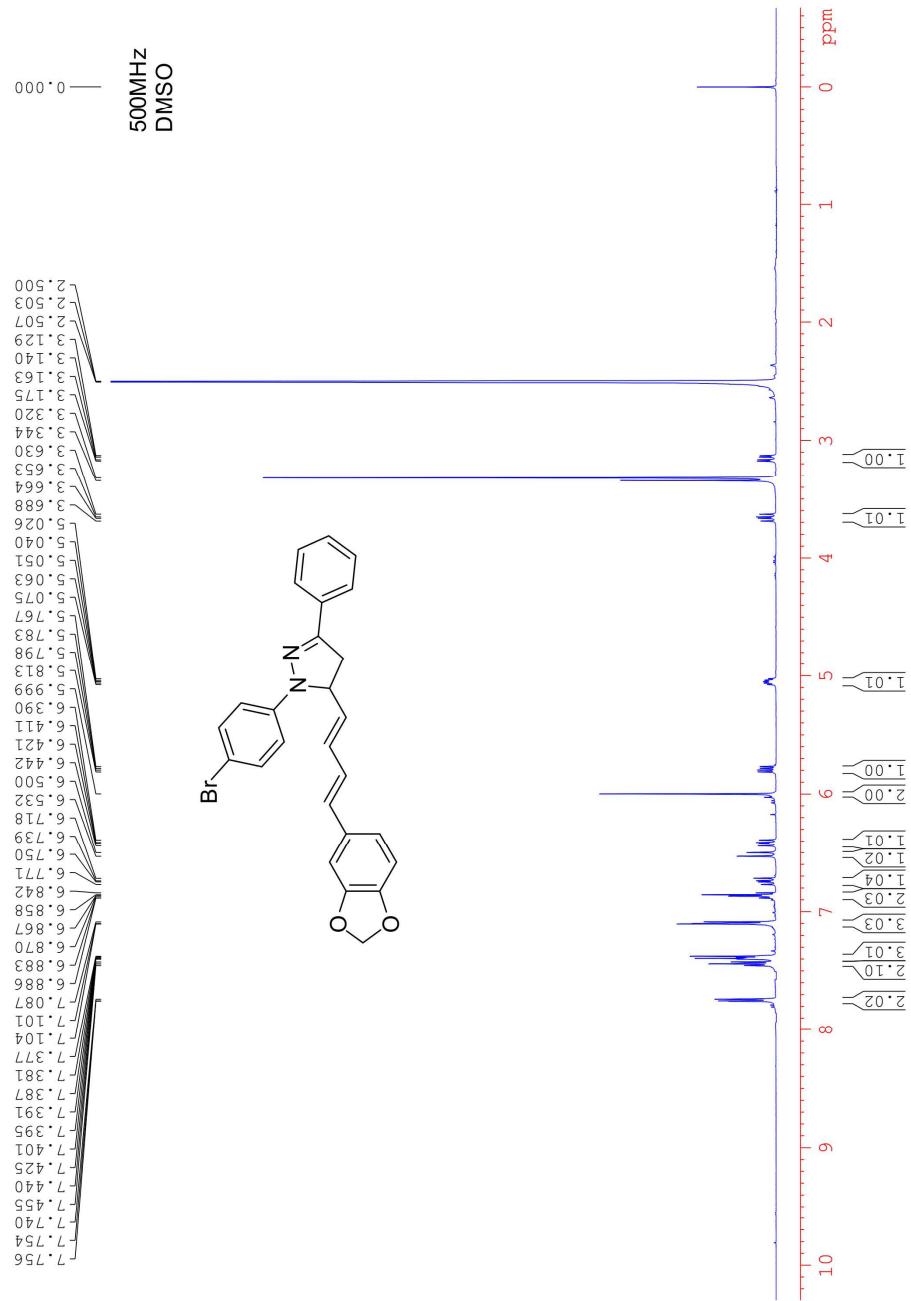
compound VIIc



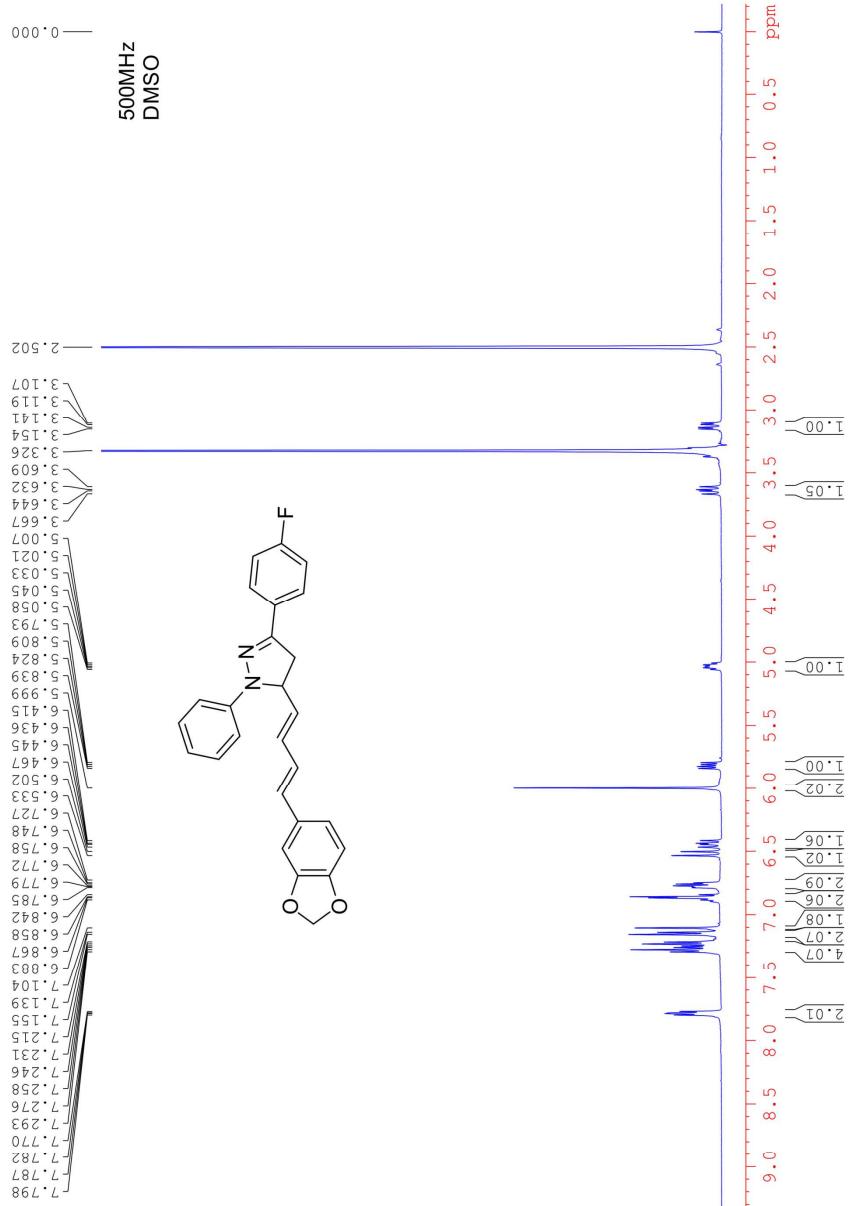
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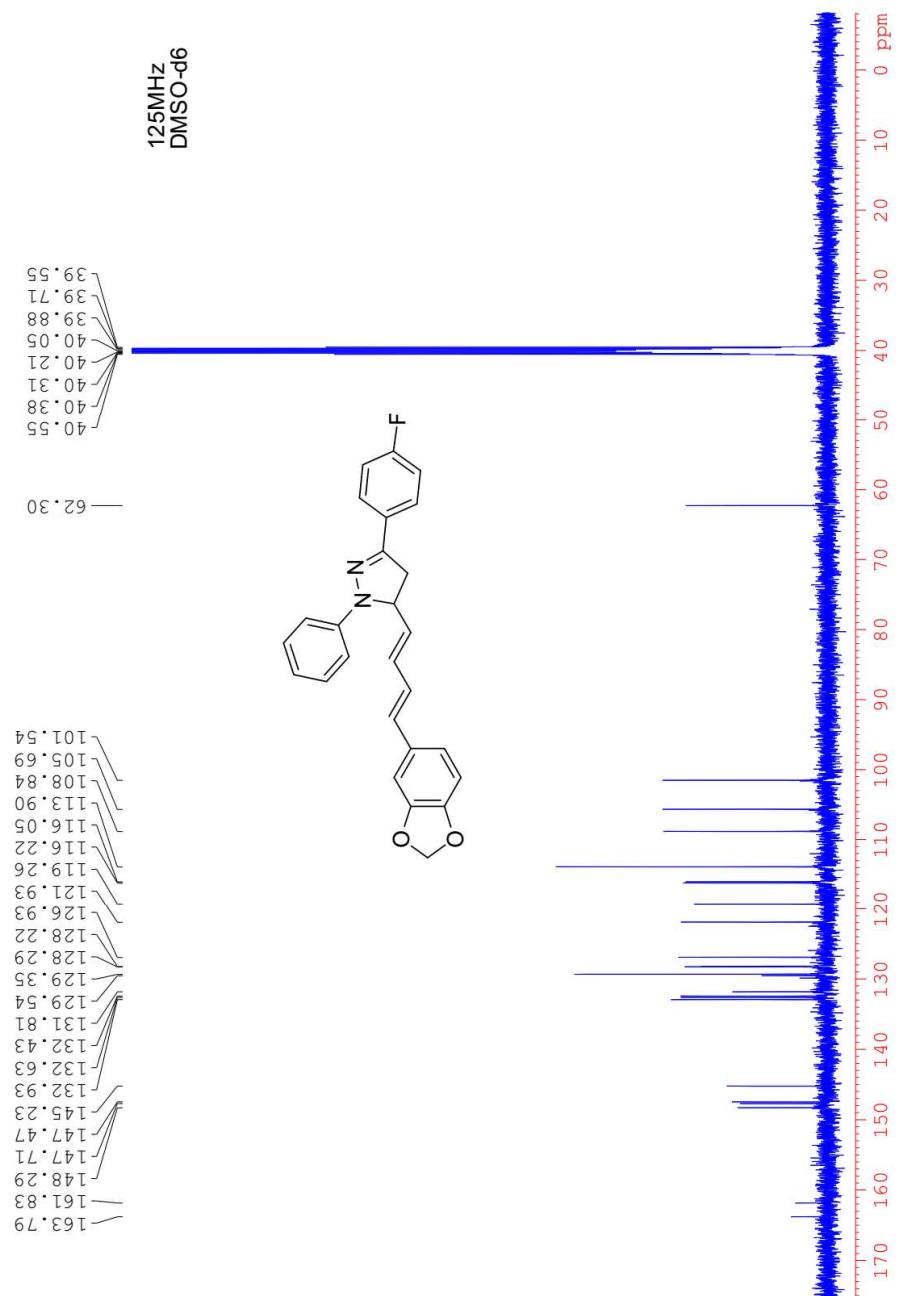
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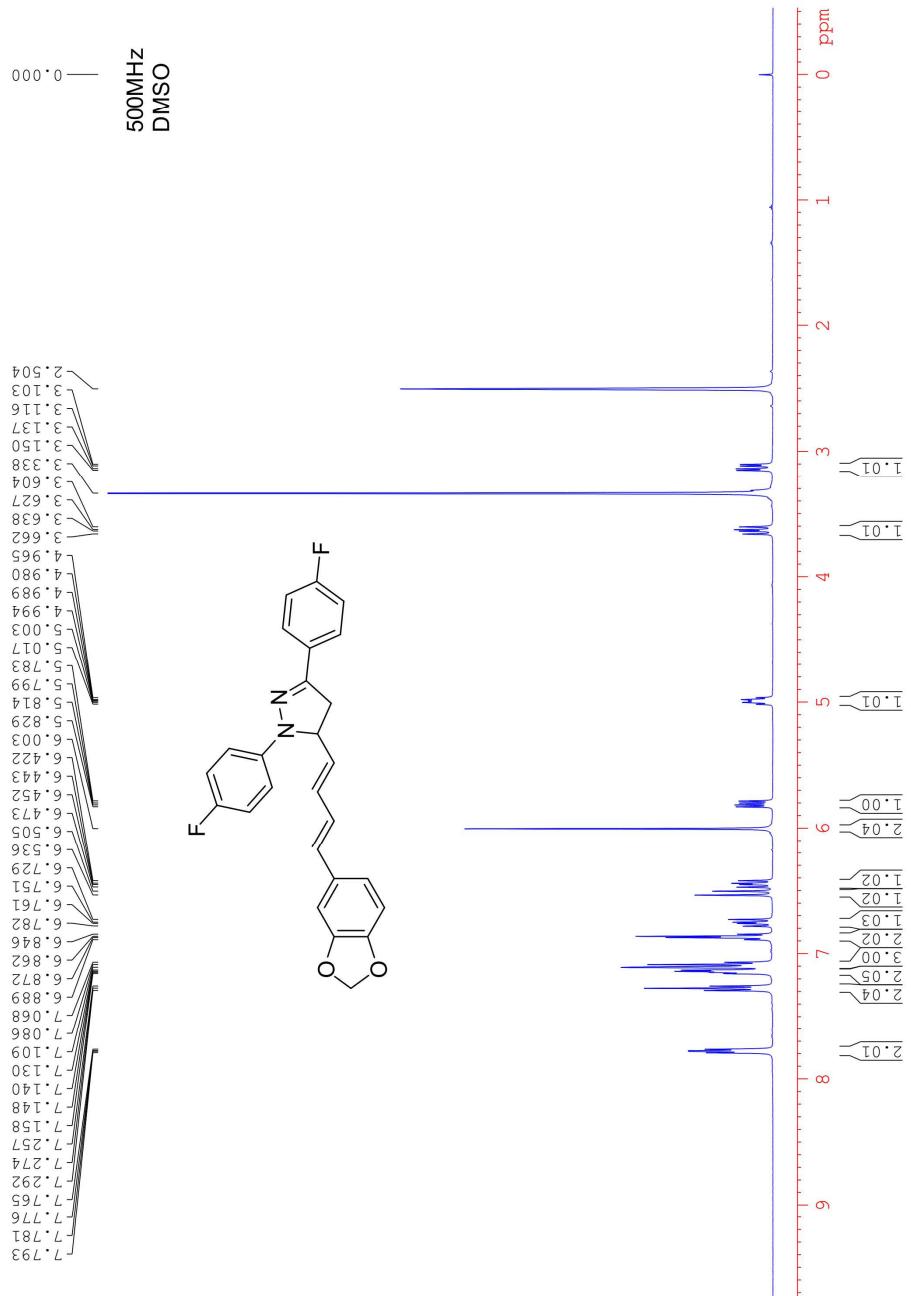
compound VIIe



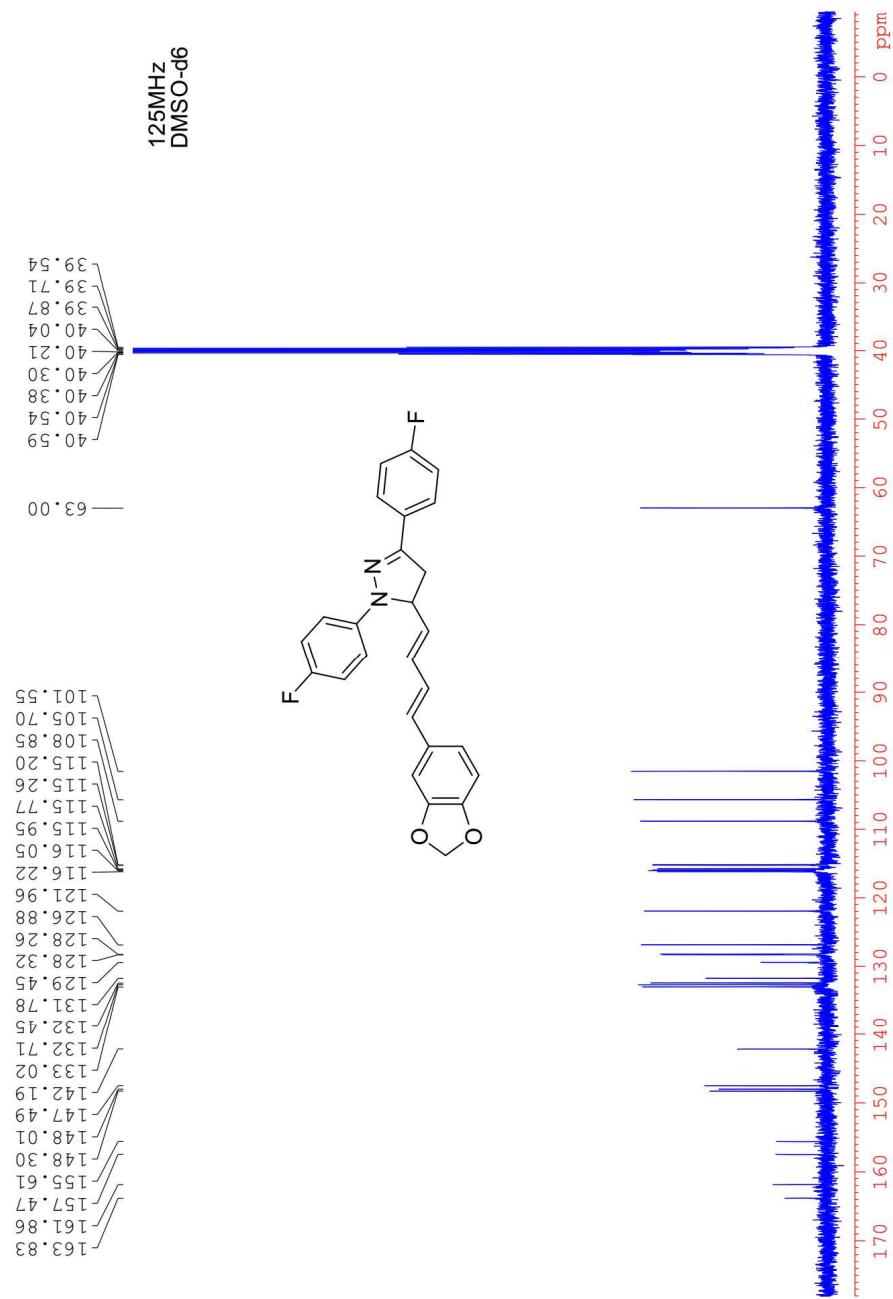
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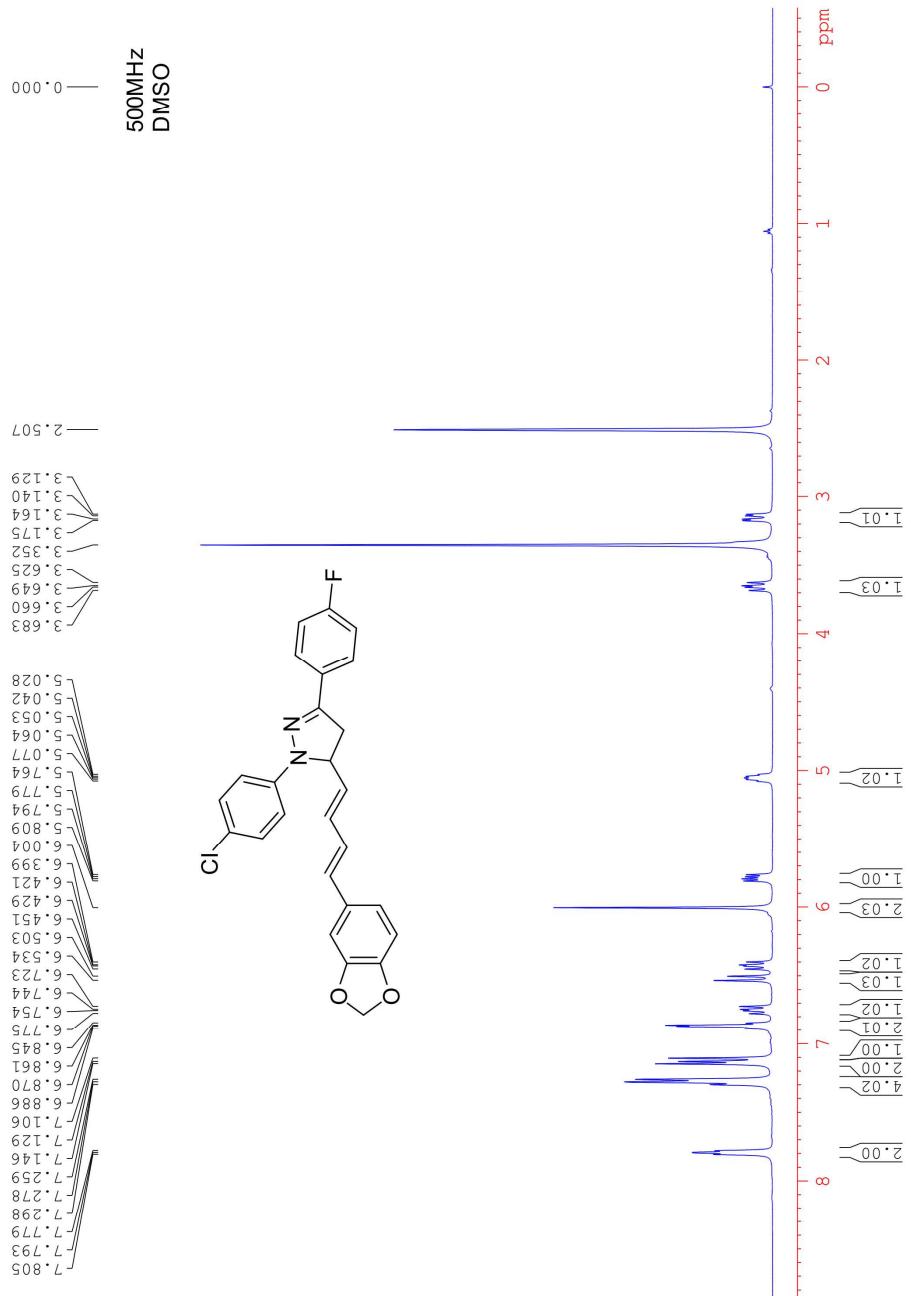
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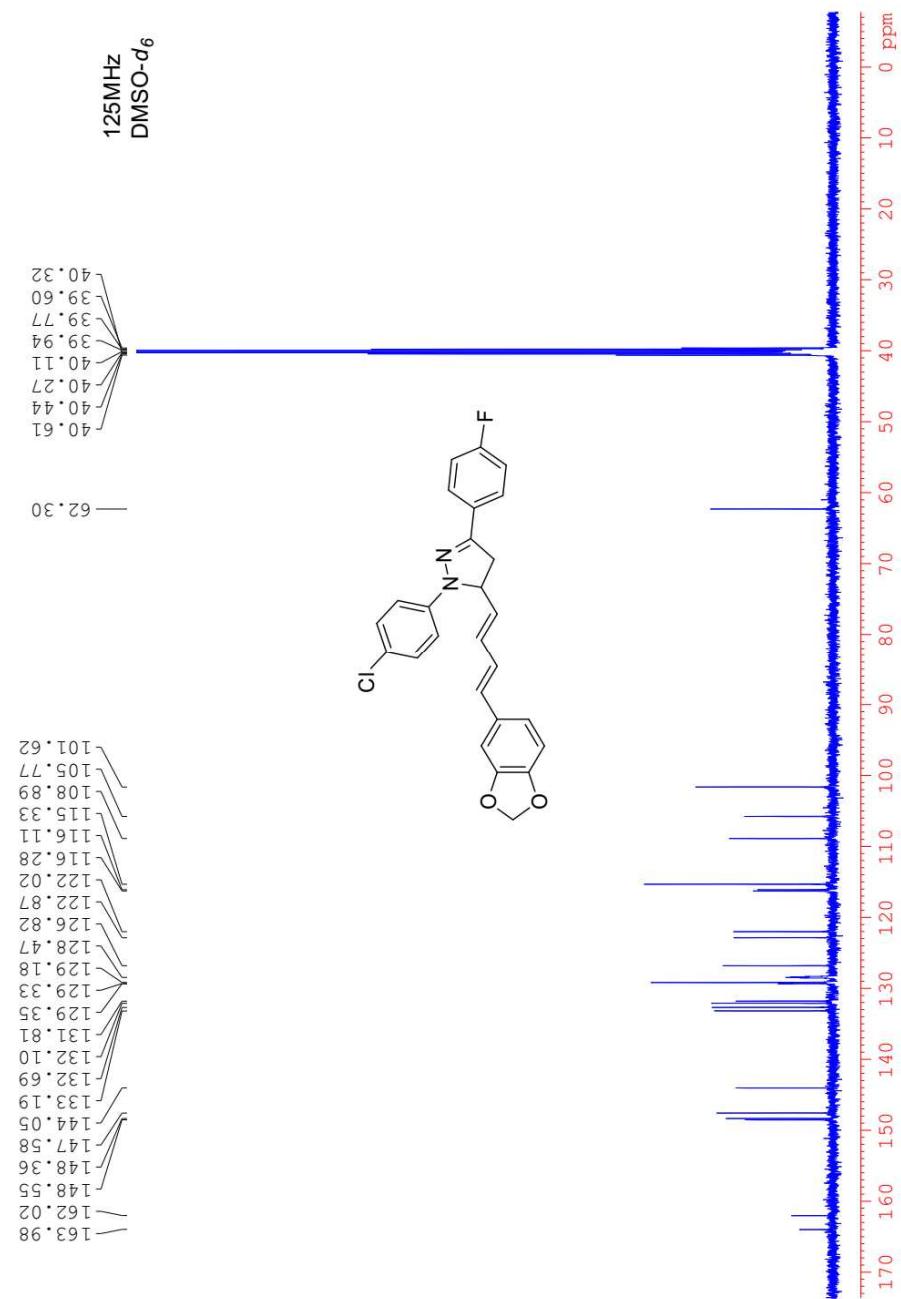
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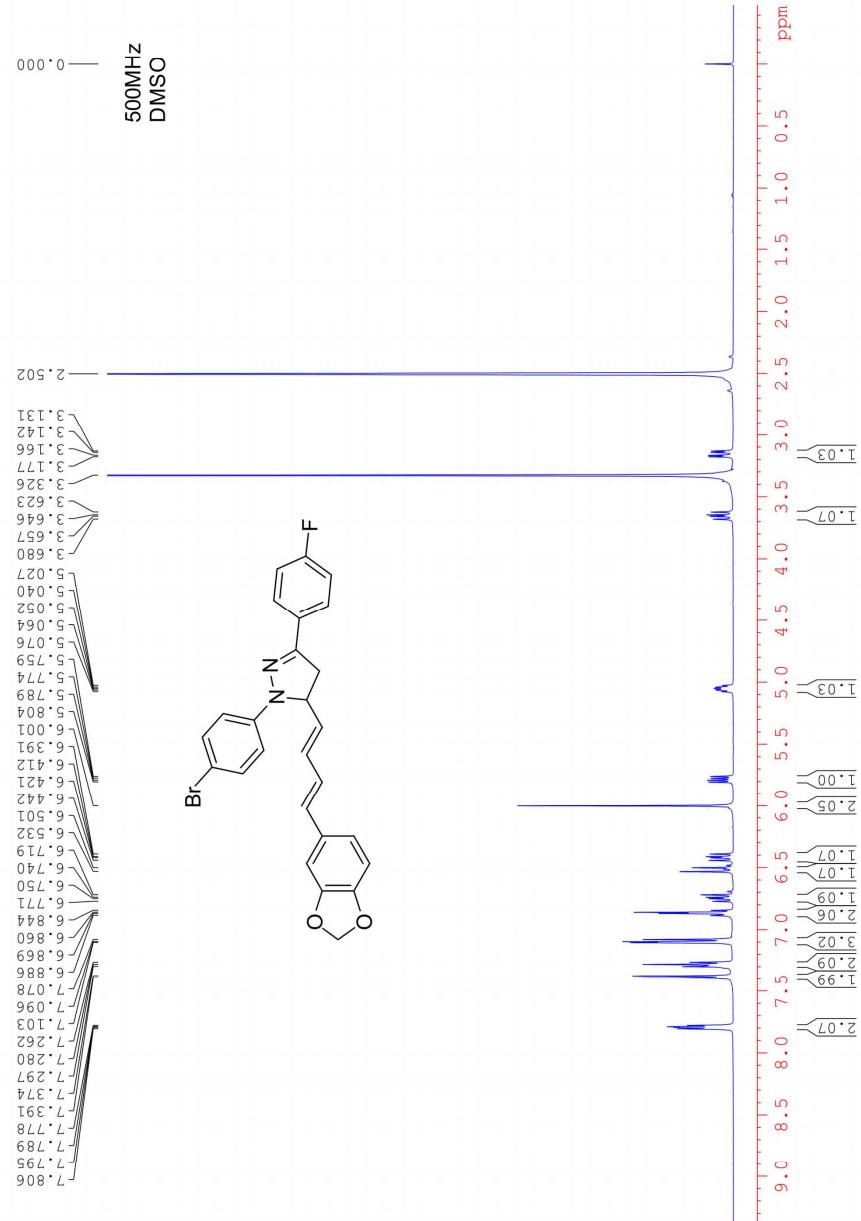
compound VIIg



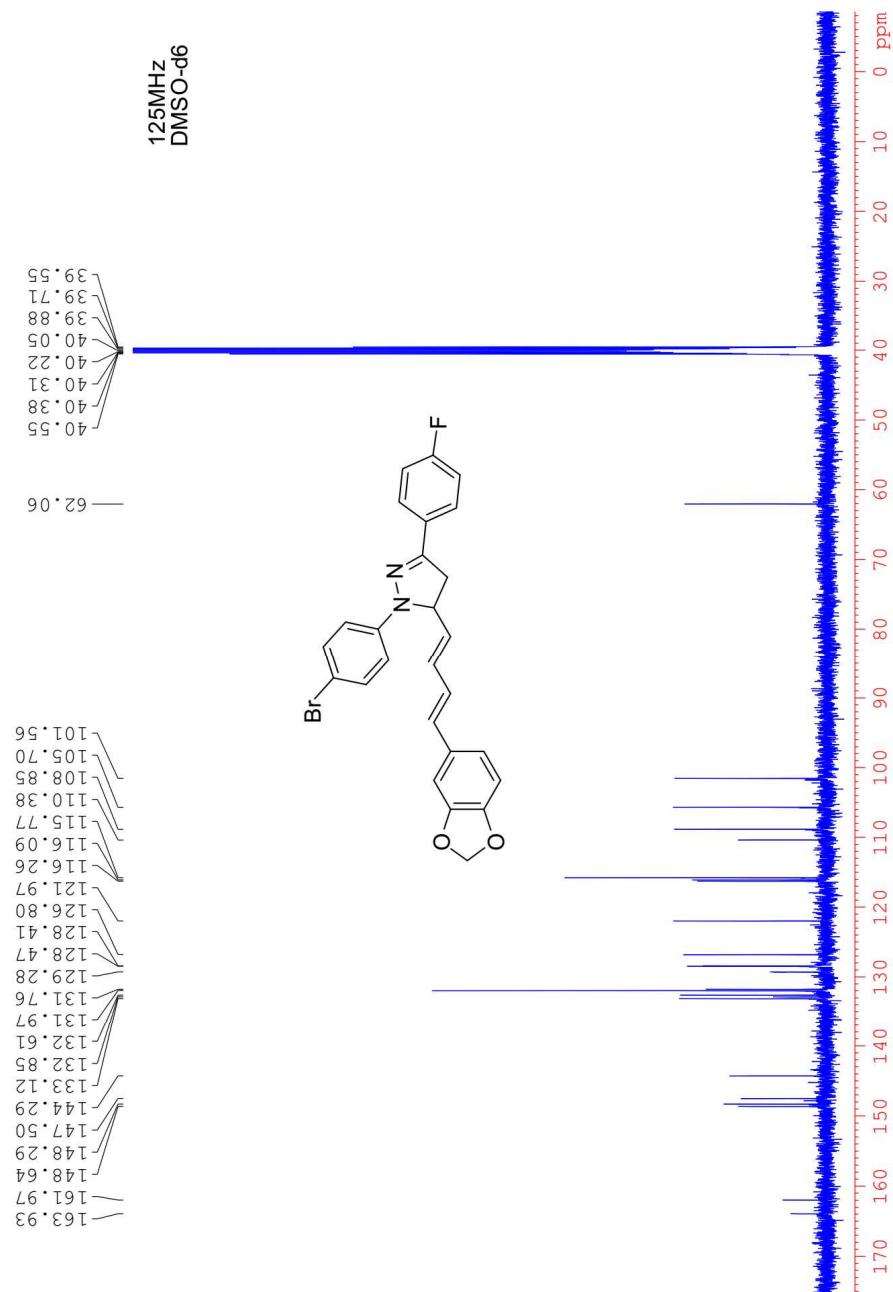
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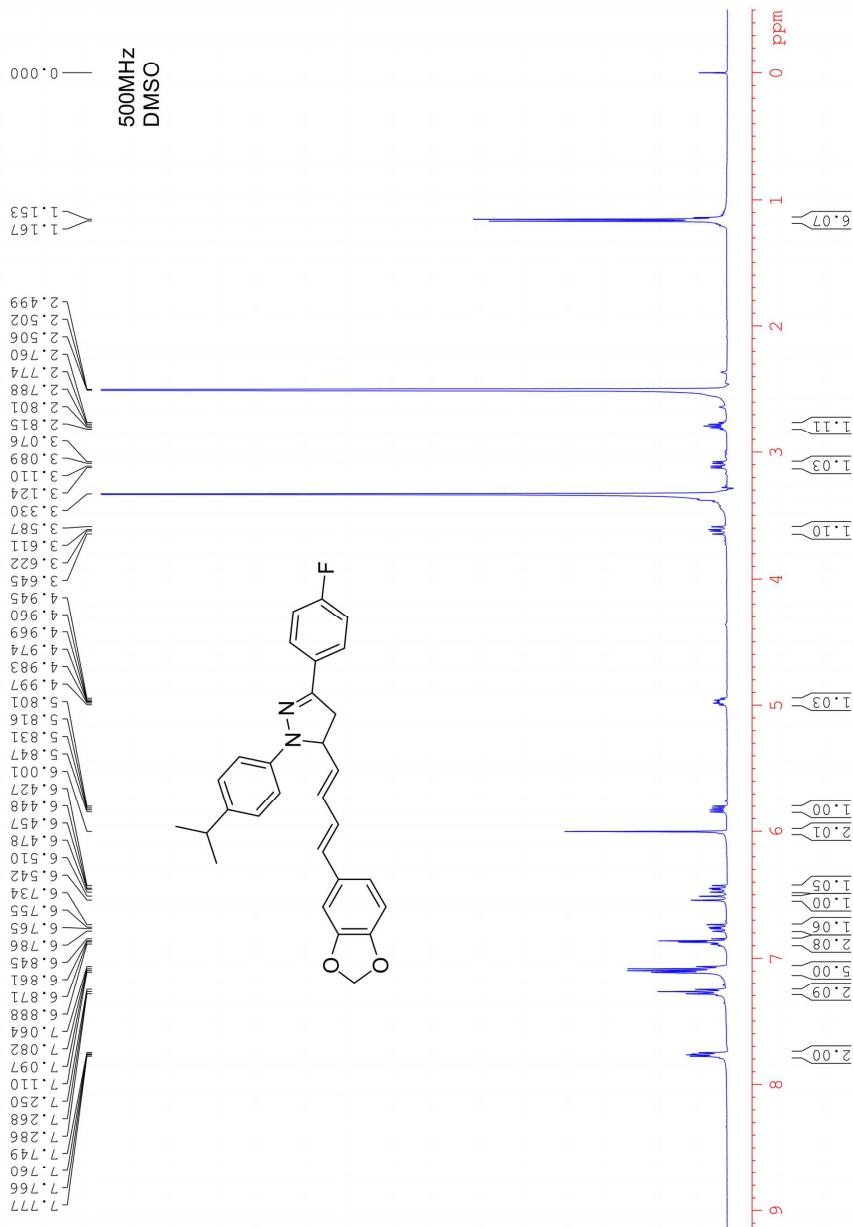
compound VIIh



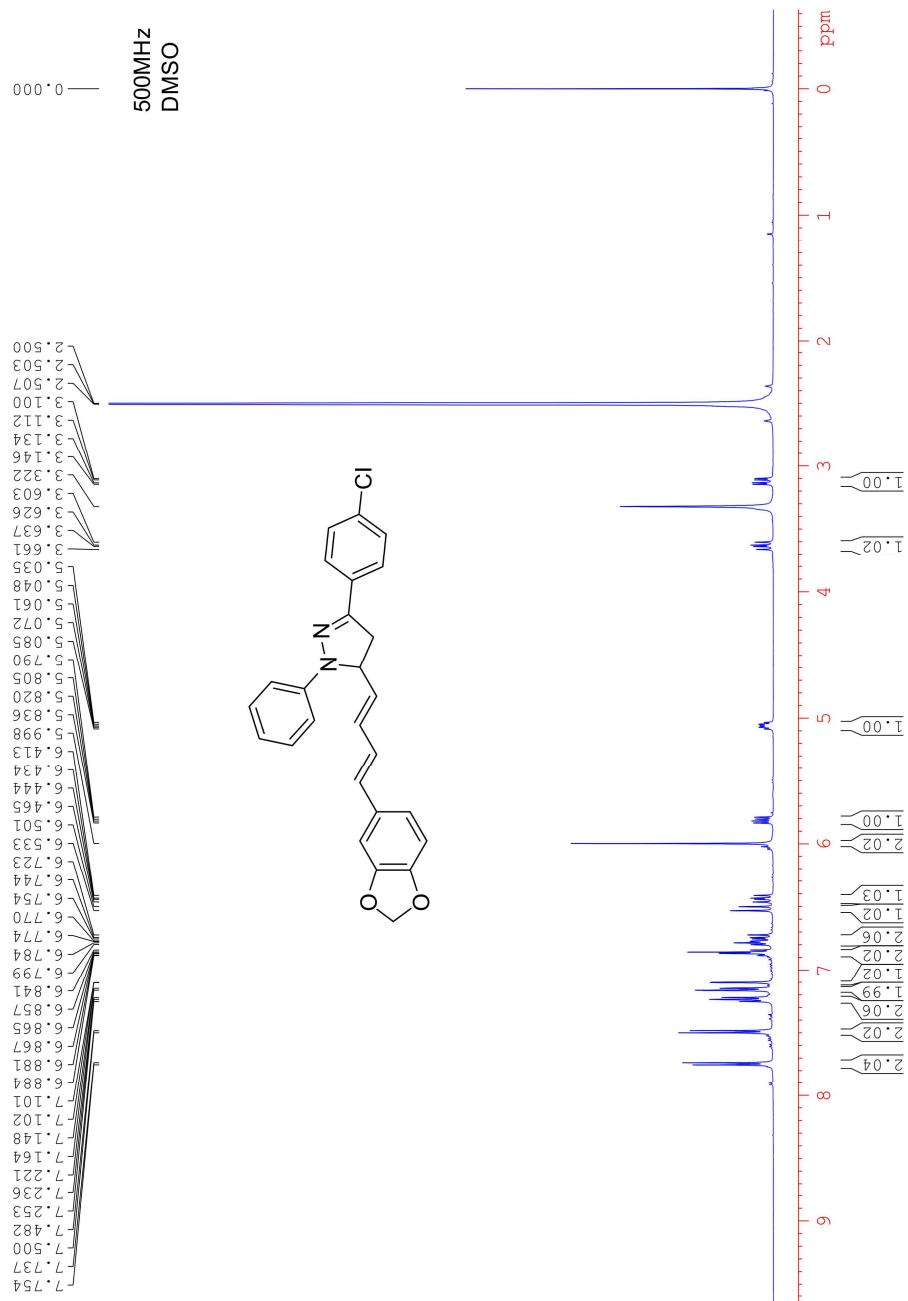
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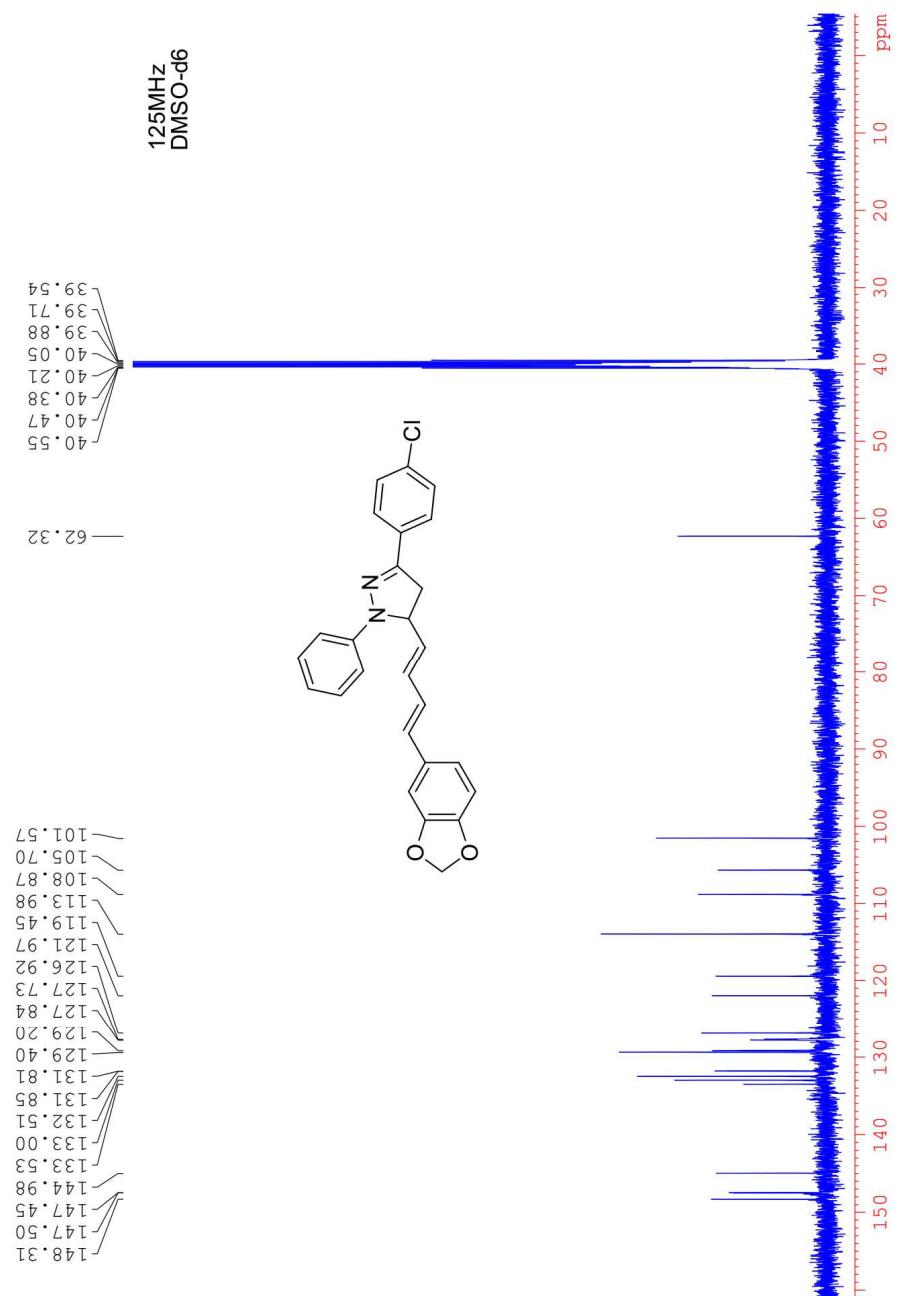
compound VIIi



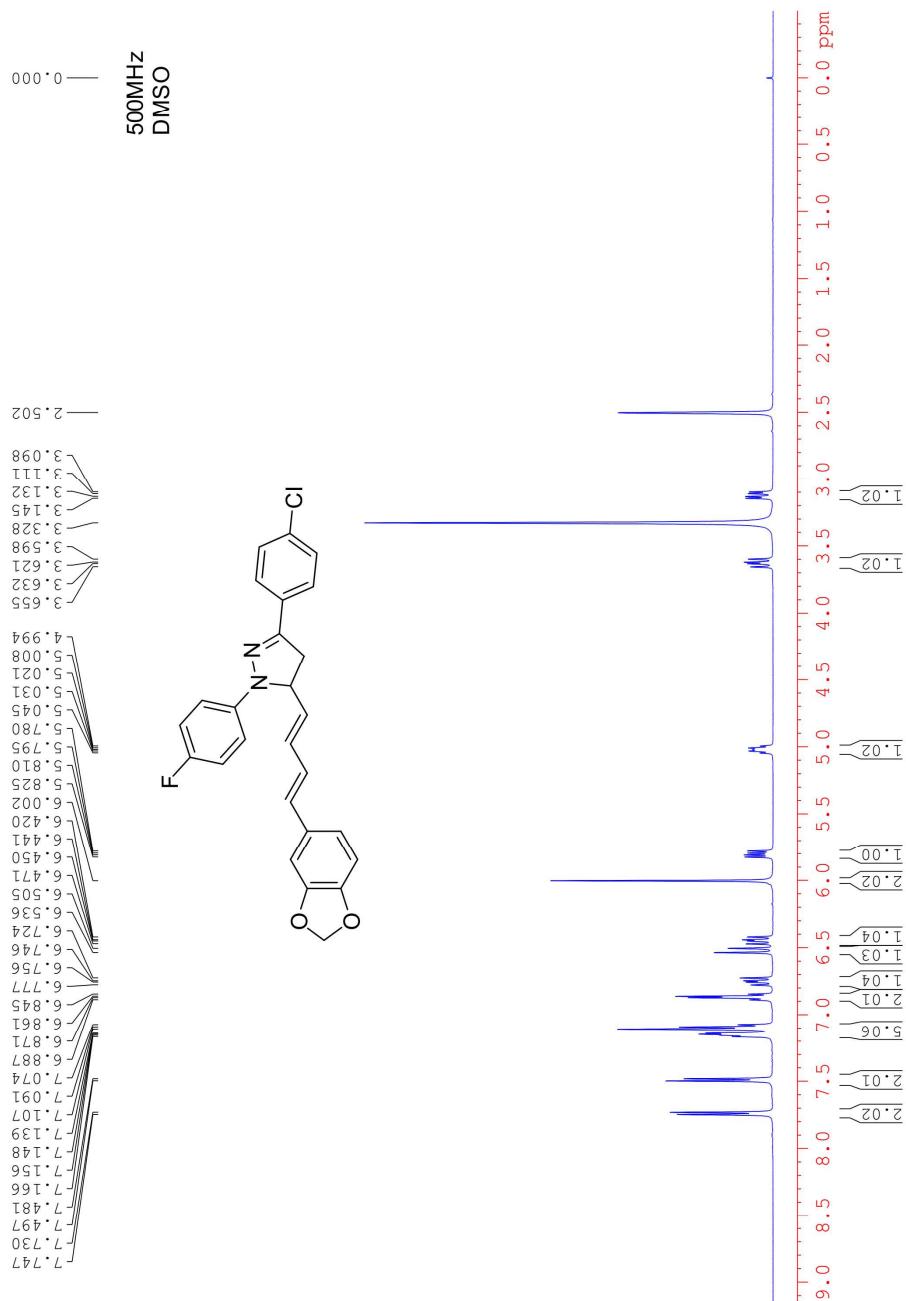
compound VIIj



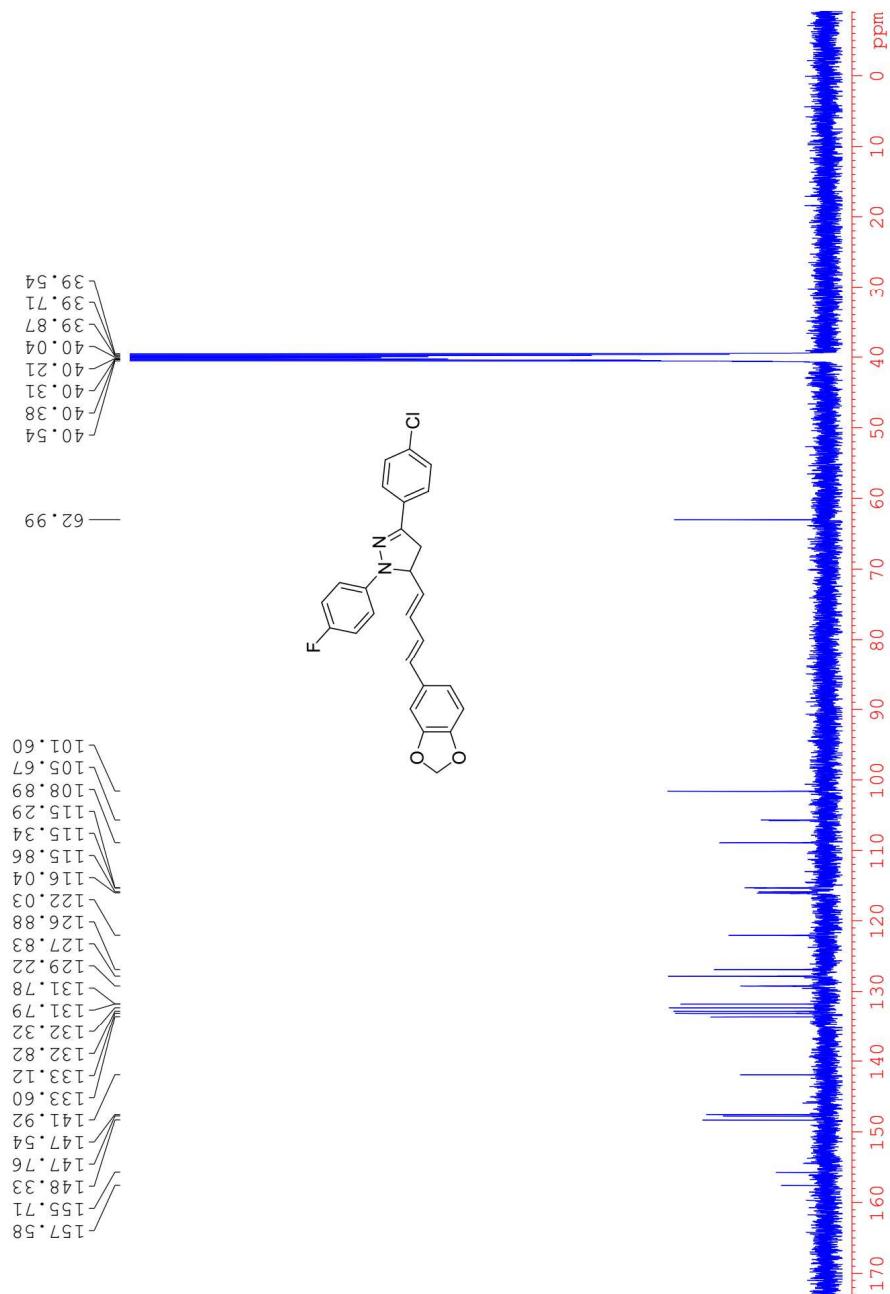
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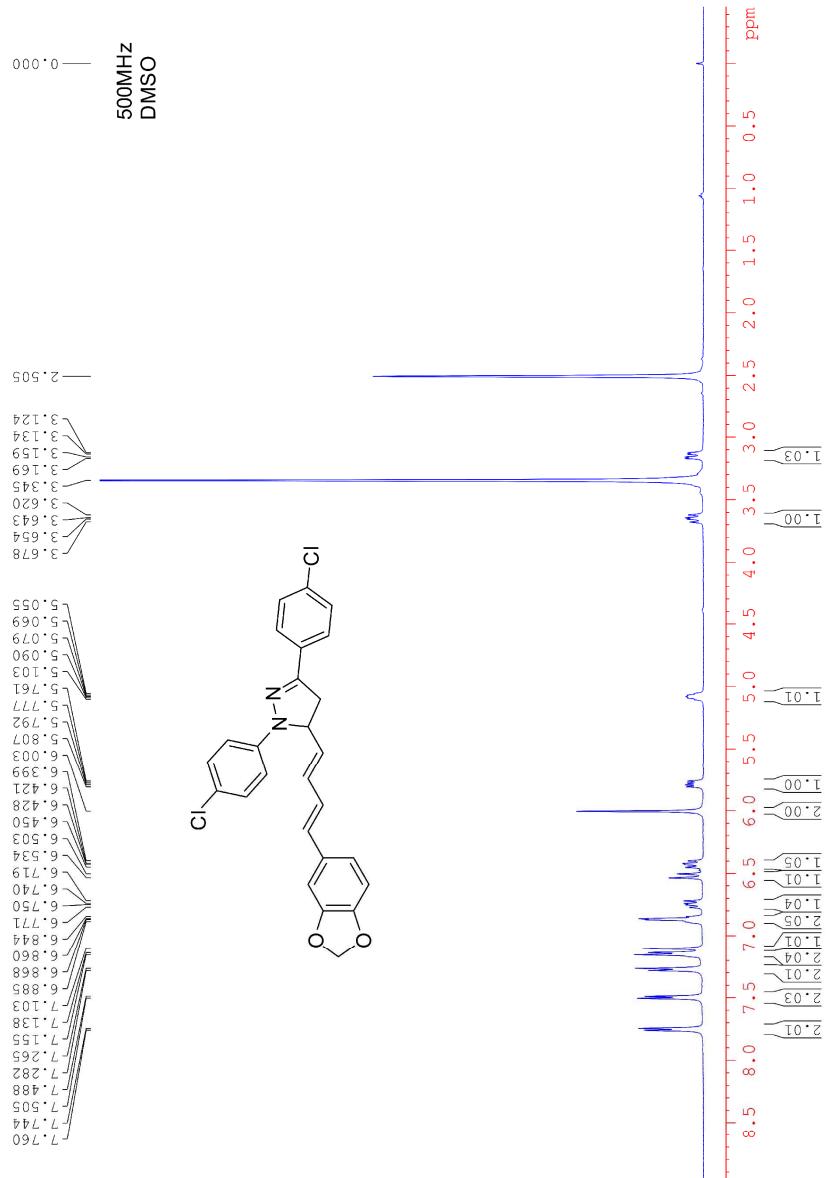
compound VIIk



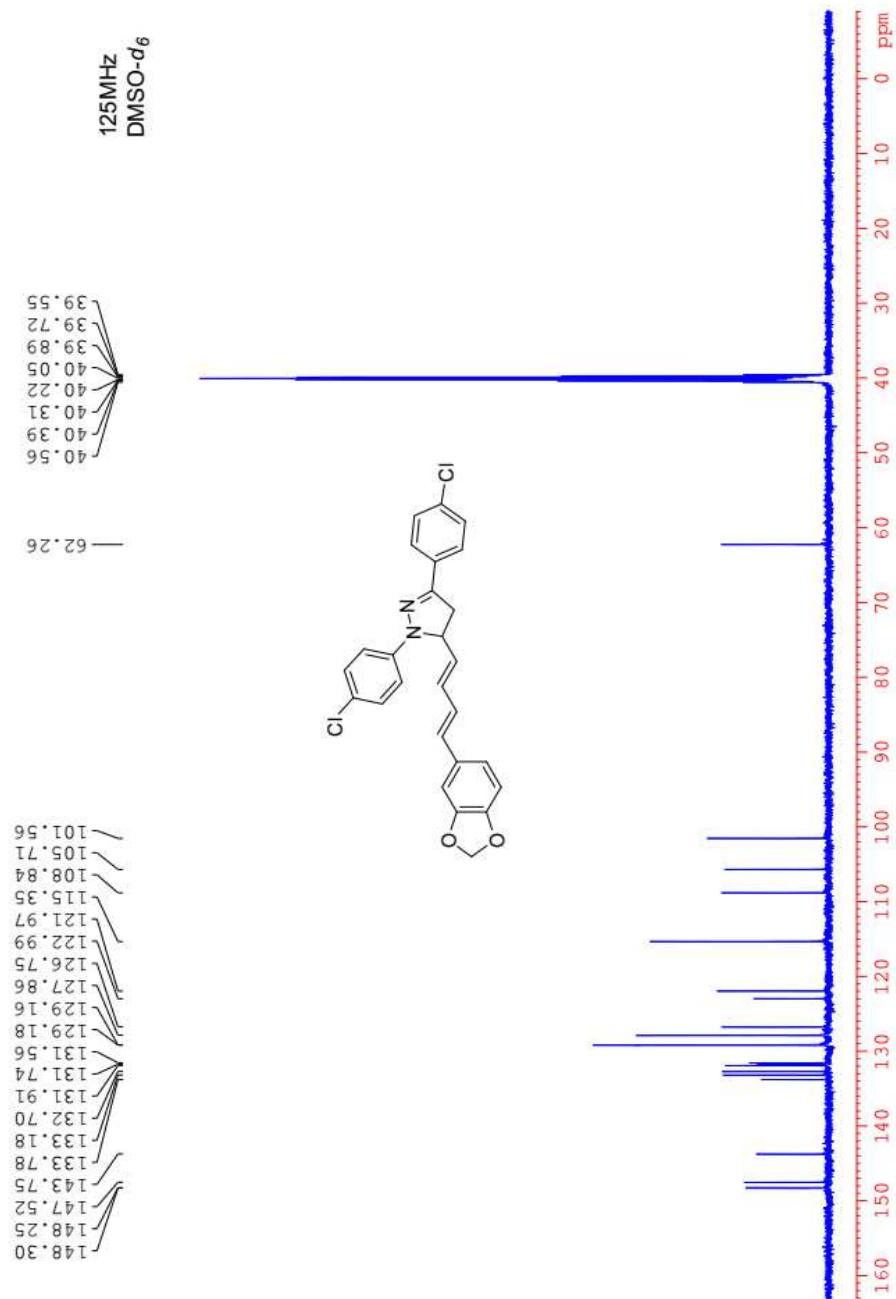
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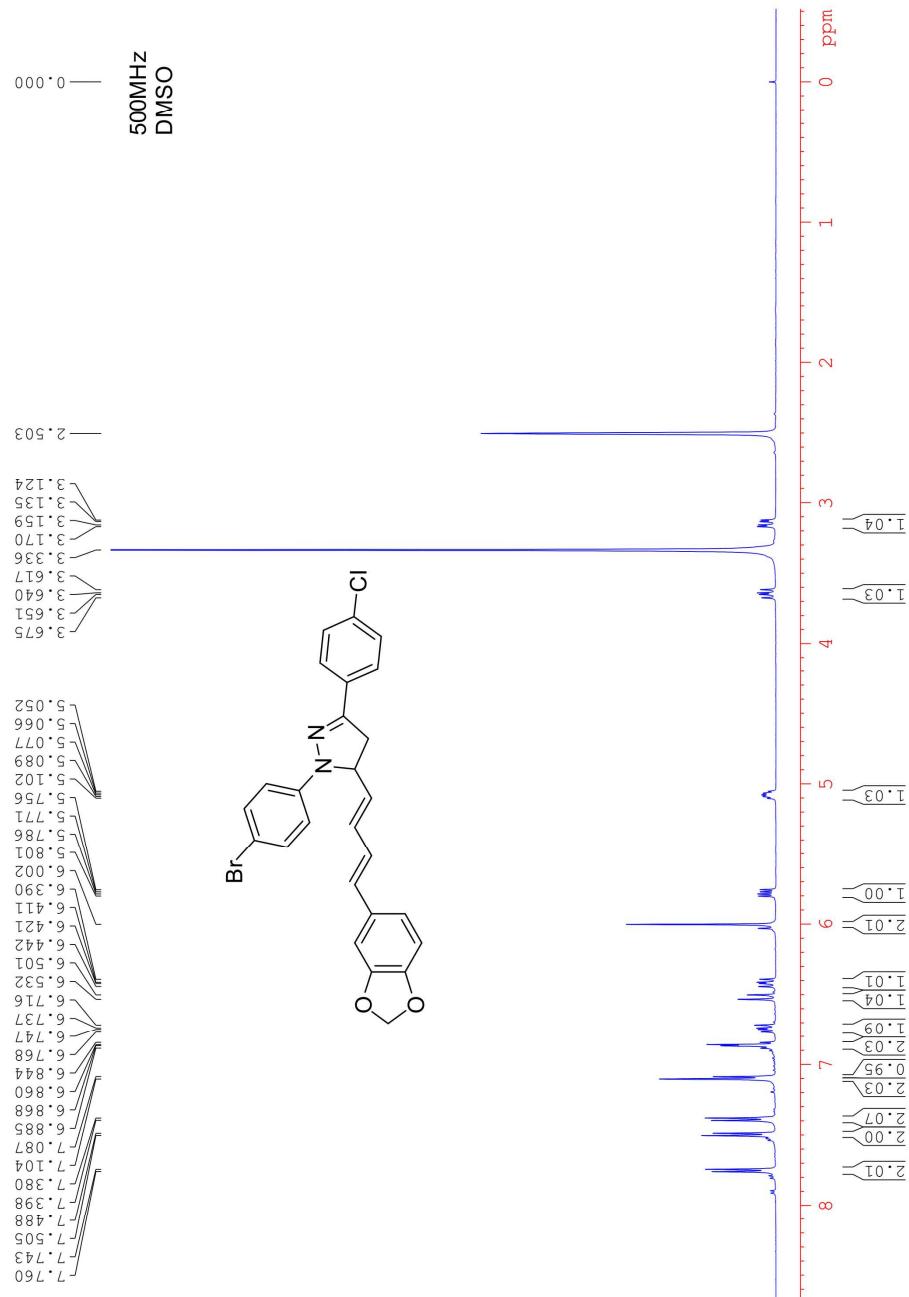
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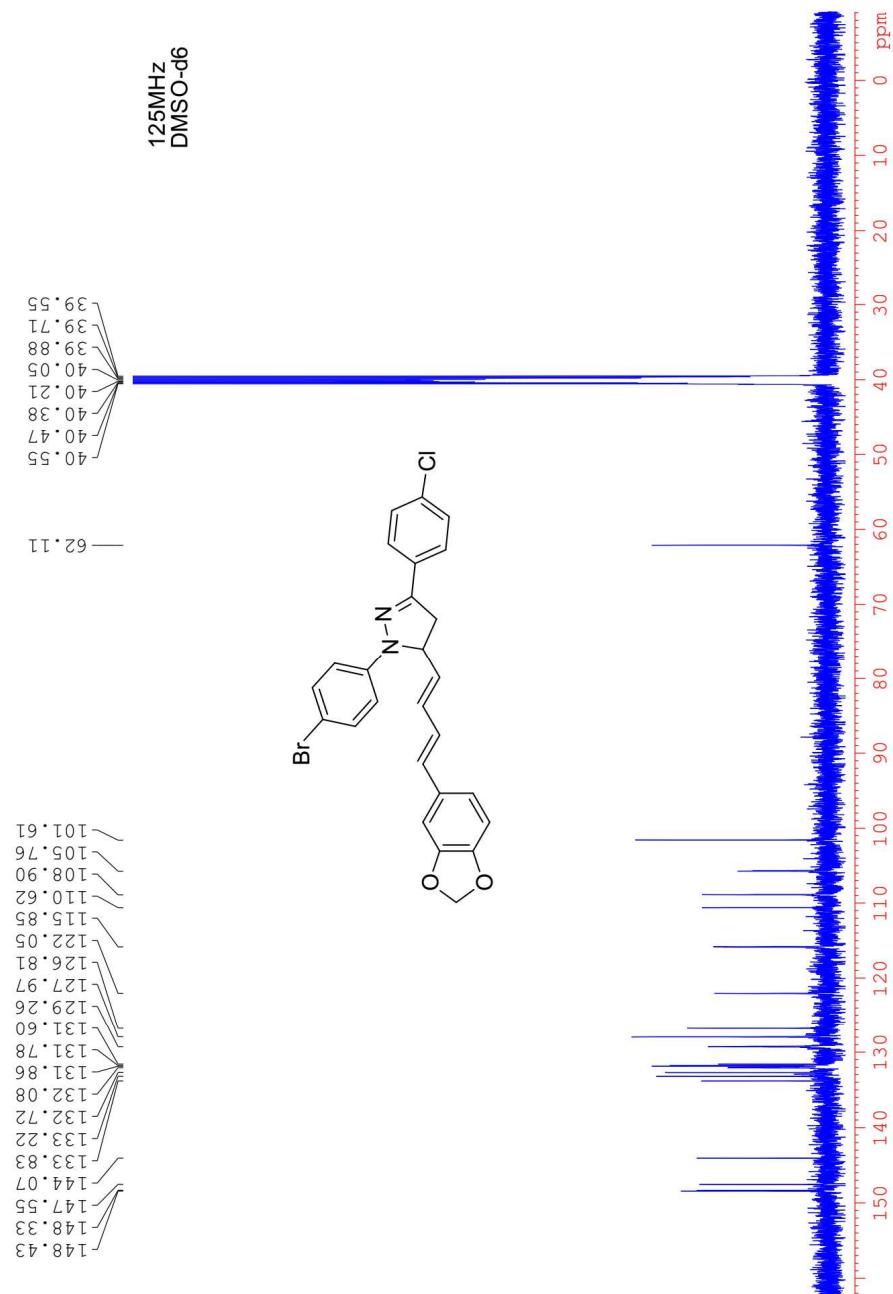
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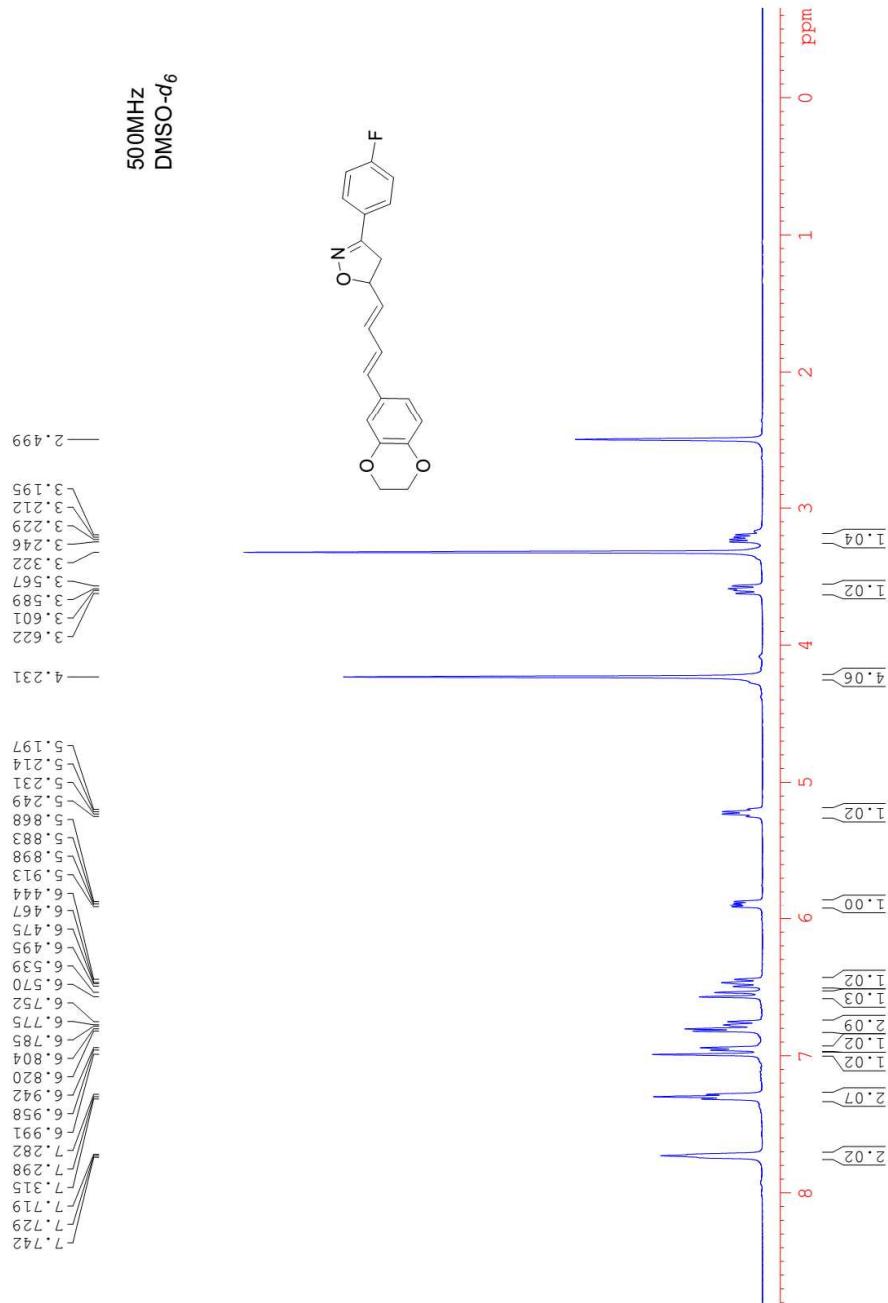
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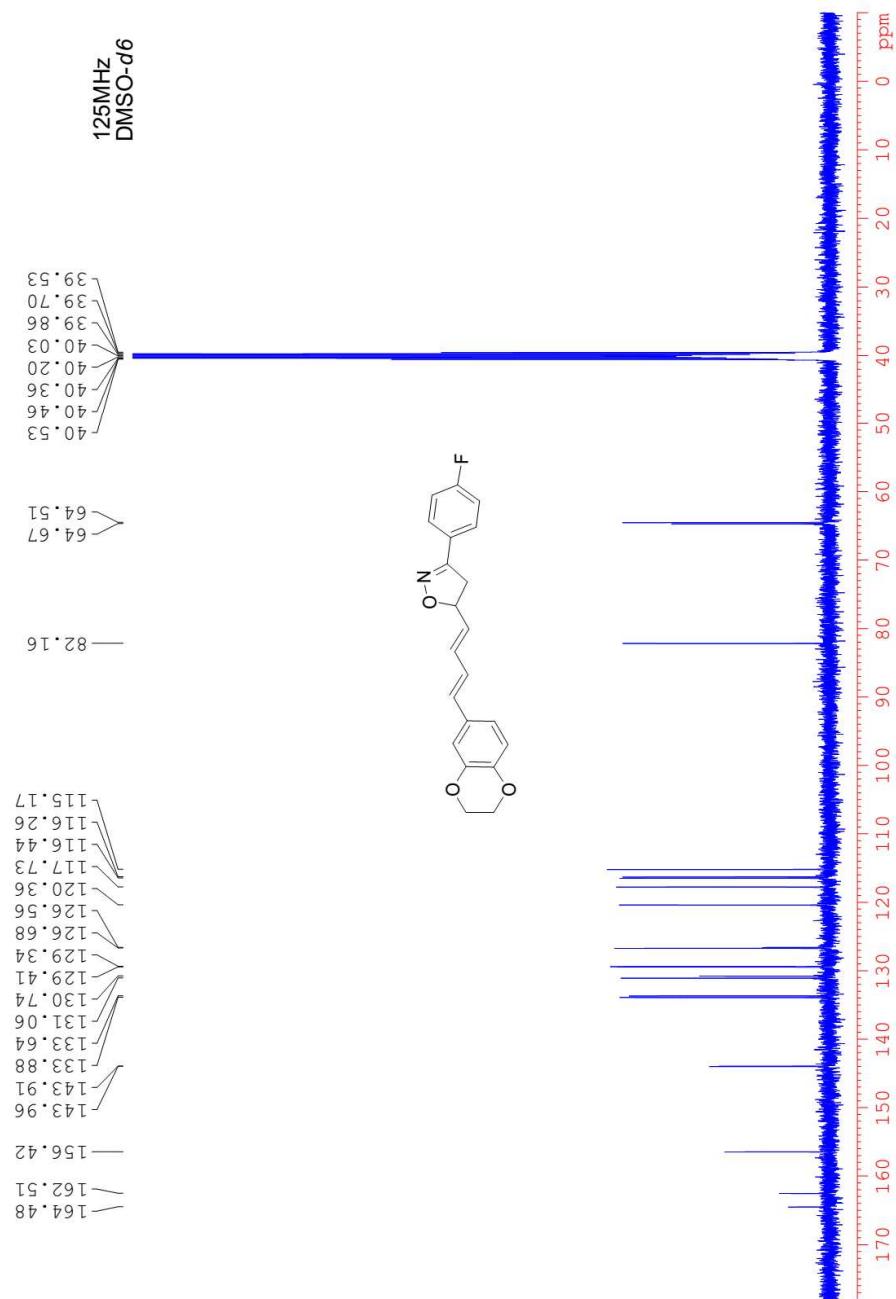
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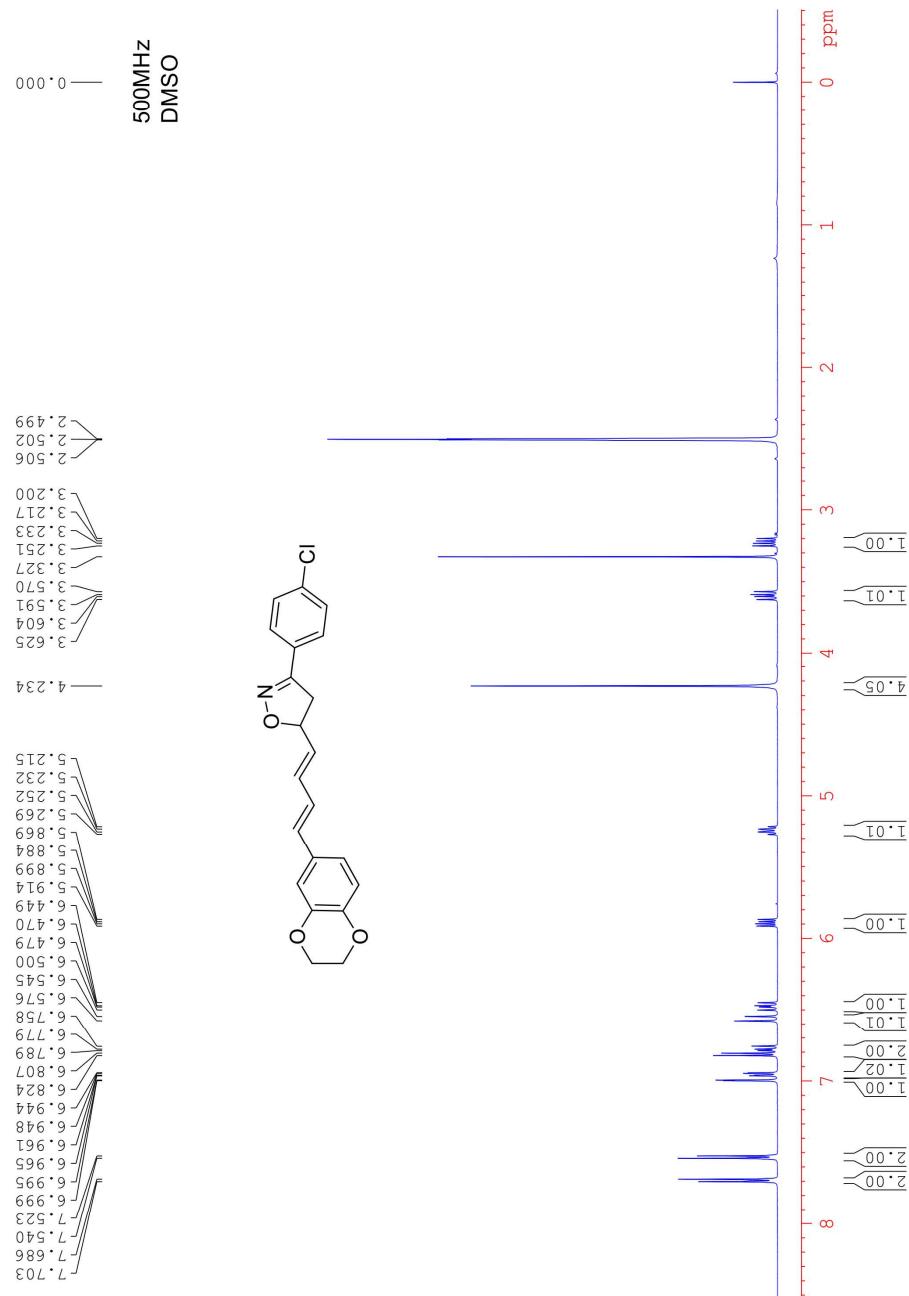
compound VIIIb



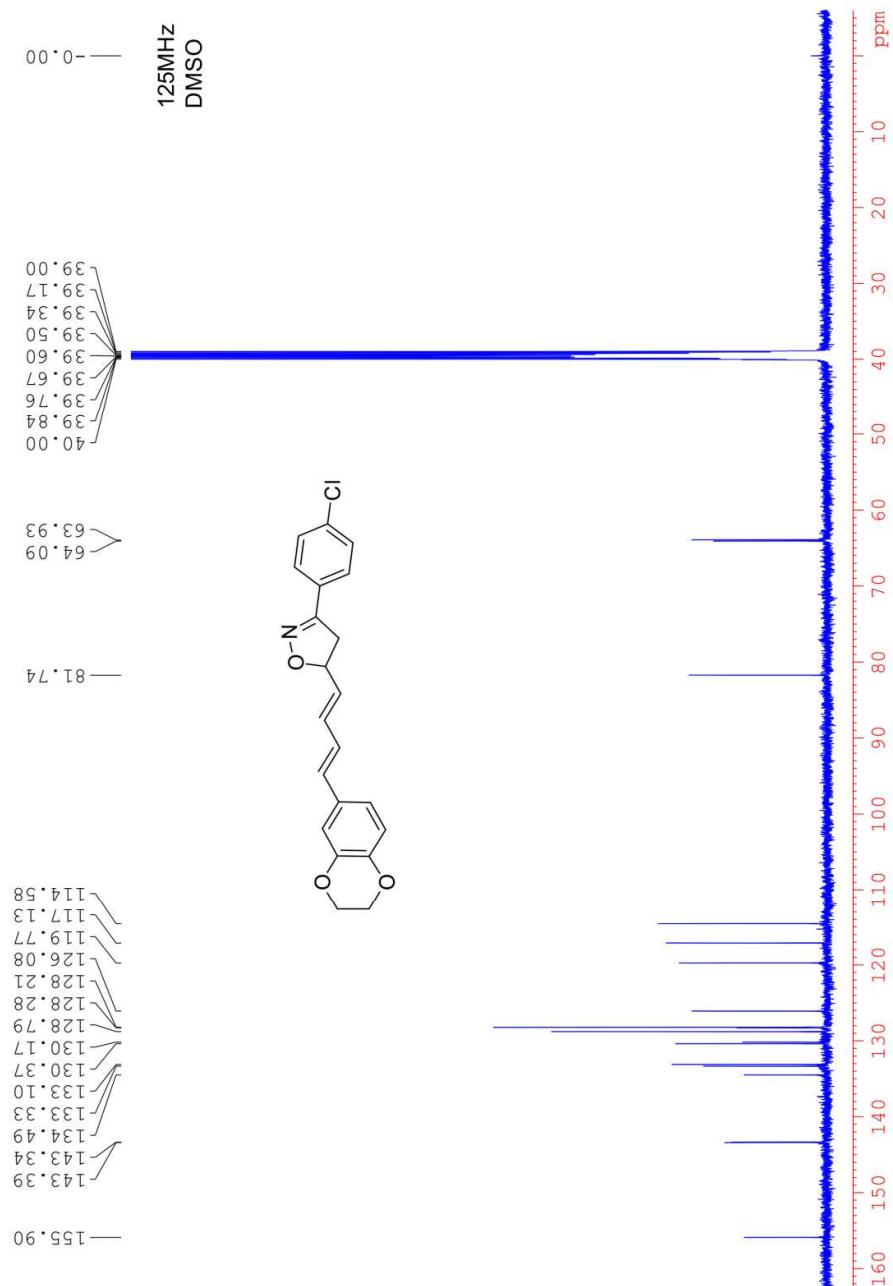
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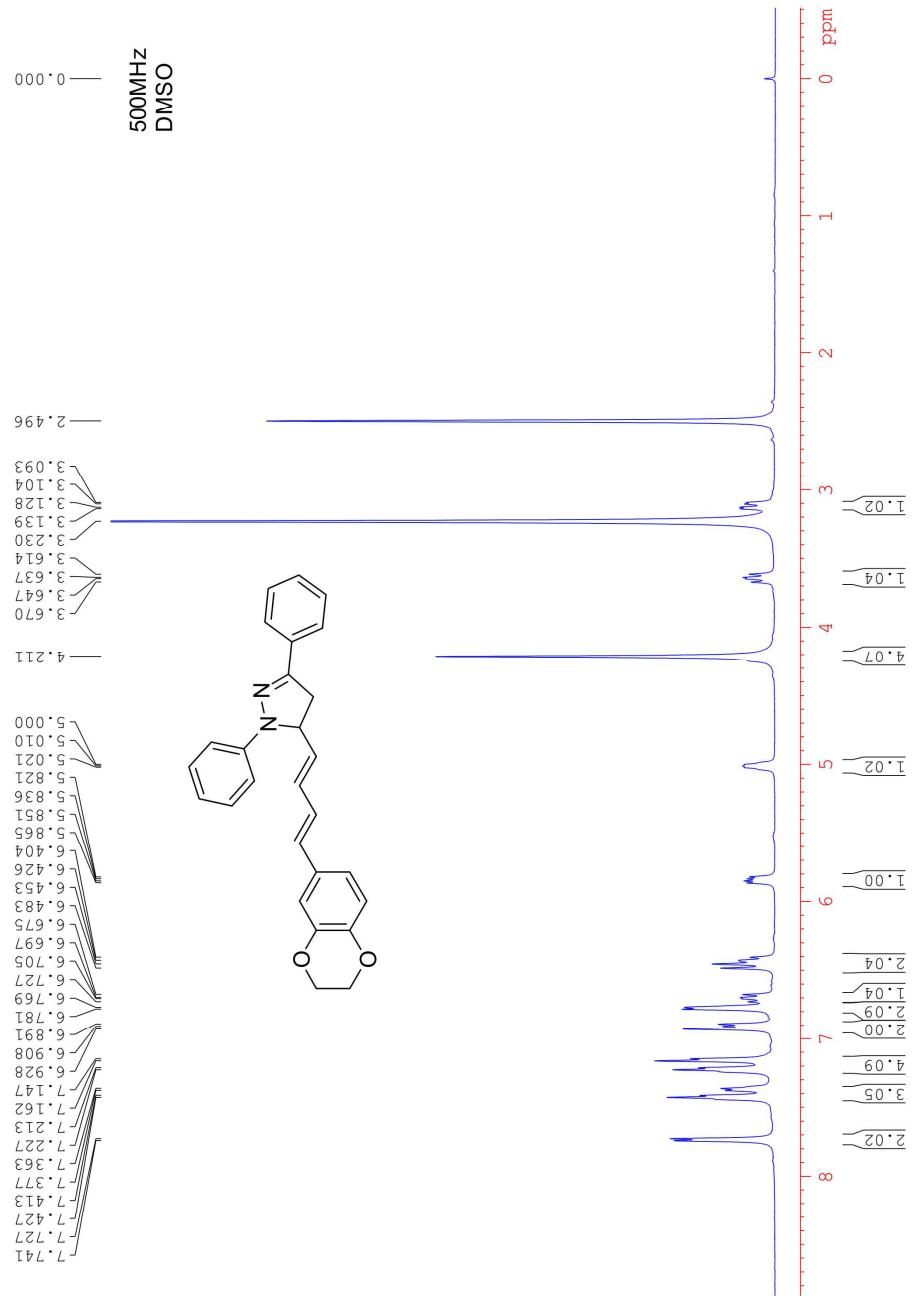
compound VIIIc



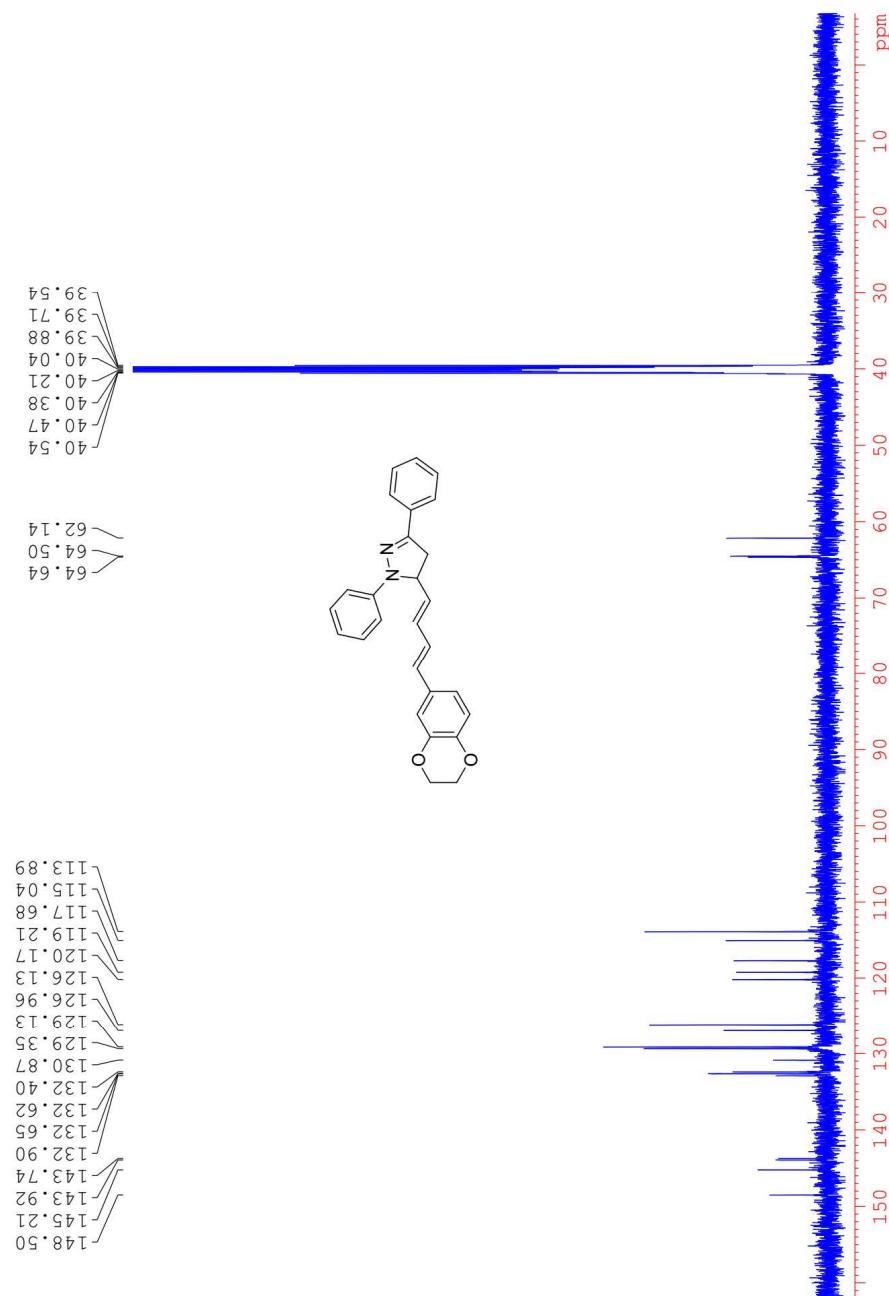
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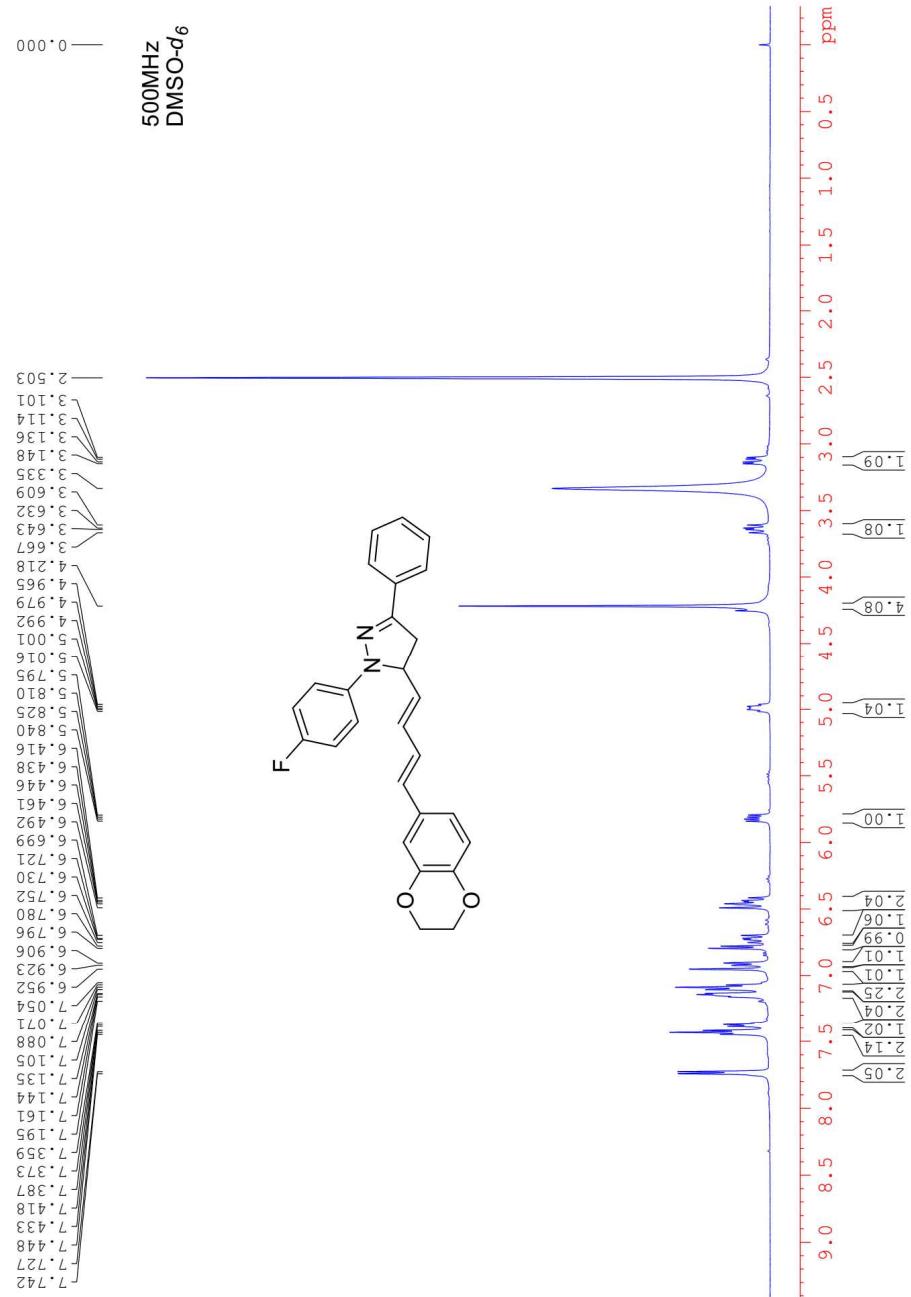
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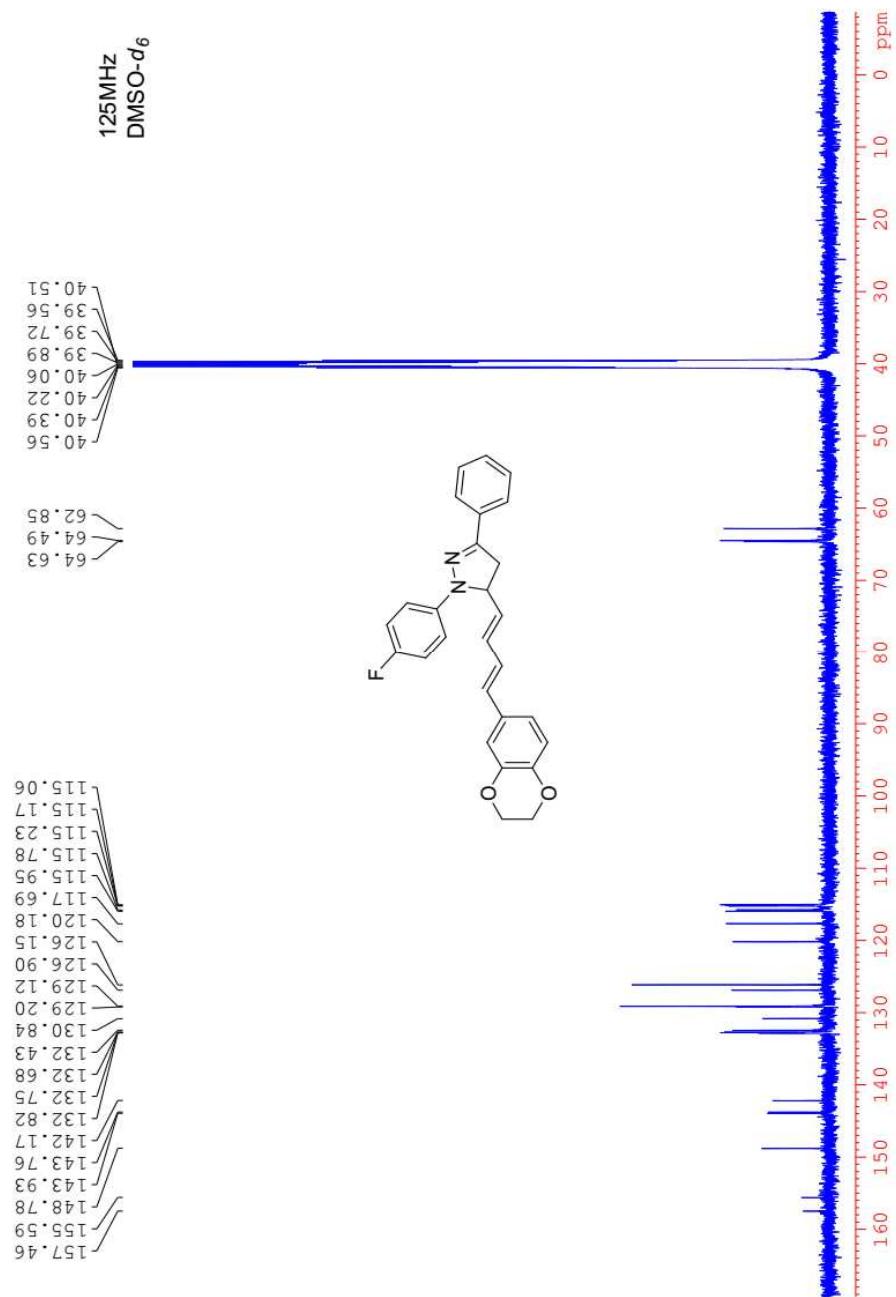
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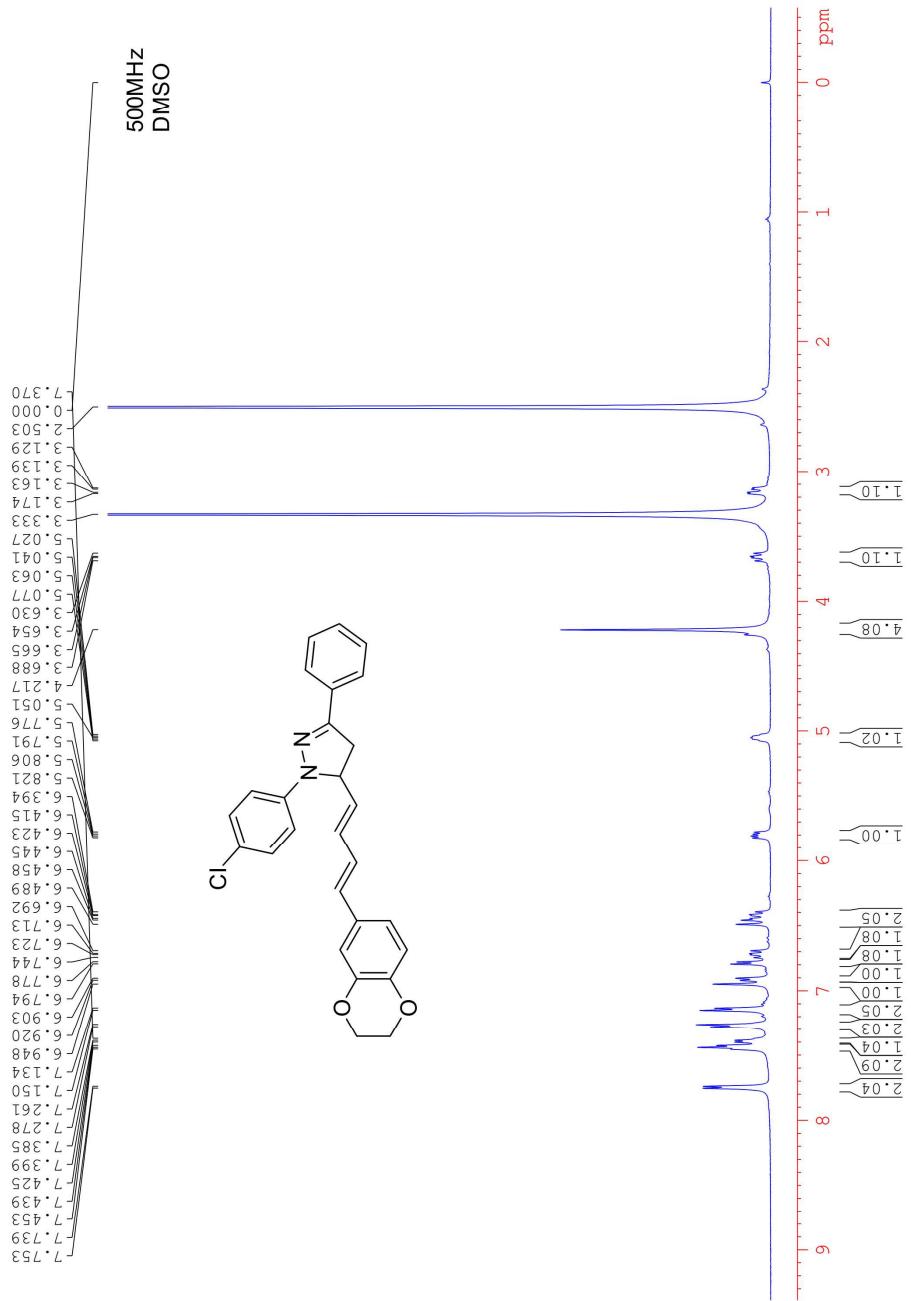
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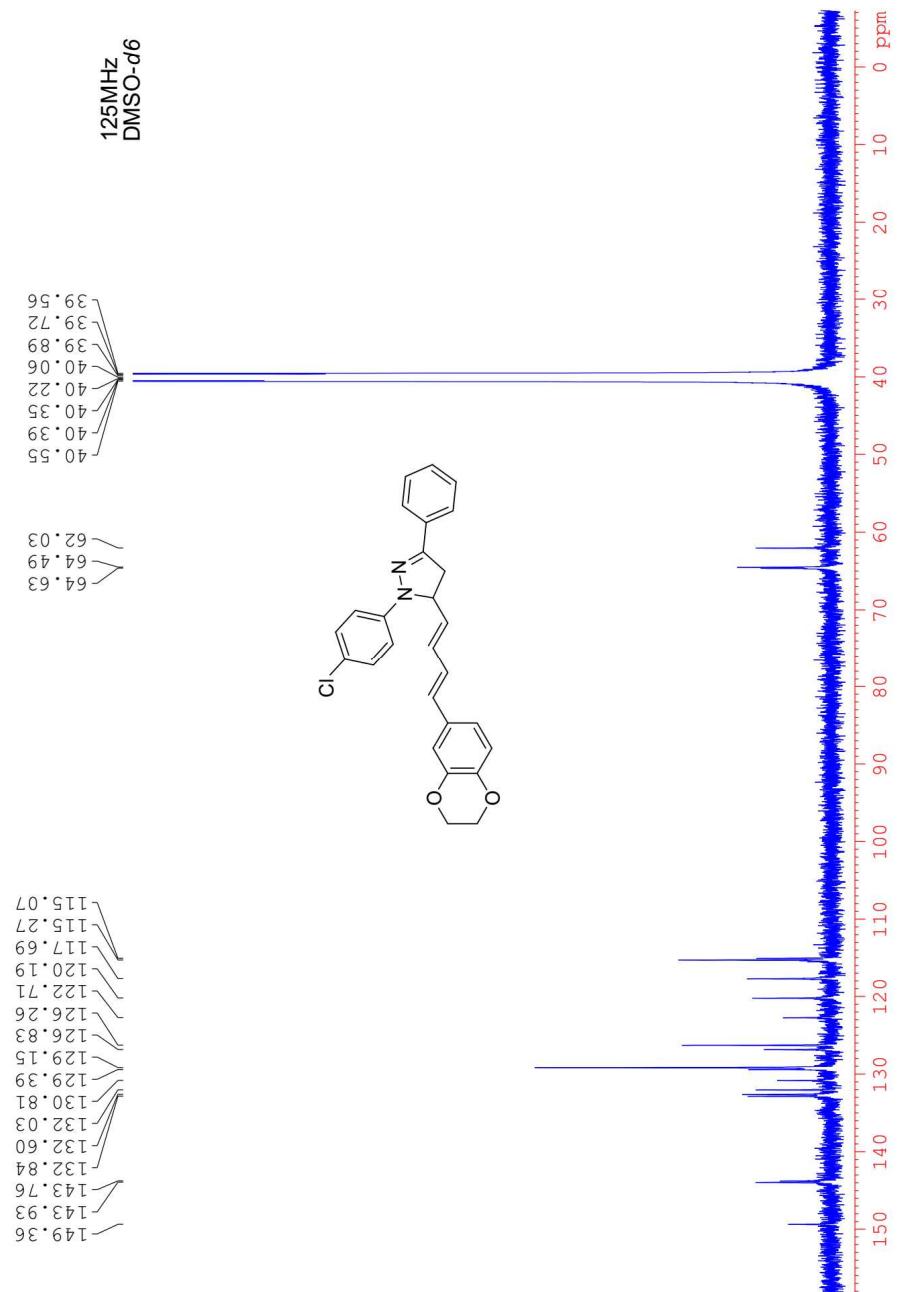
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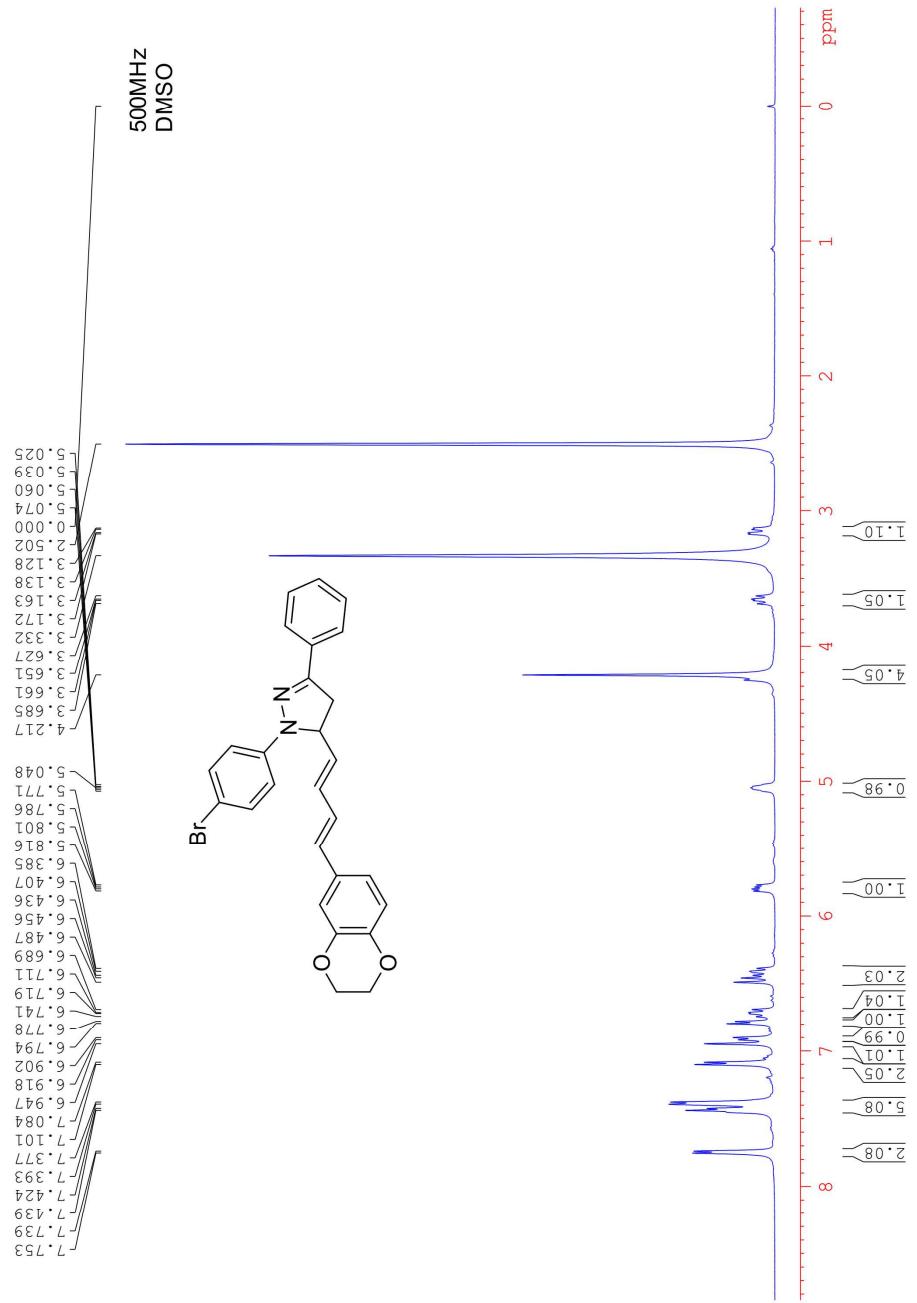
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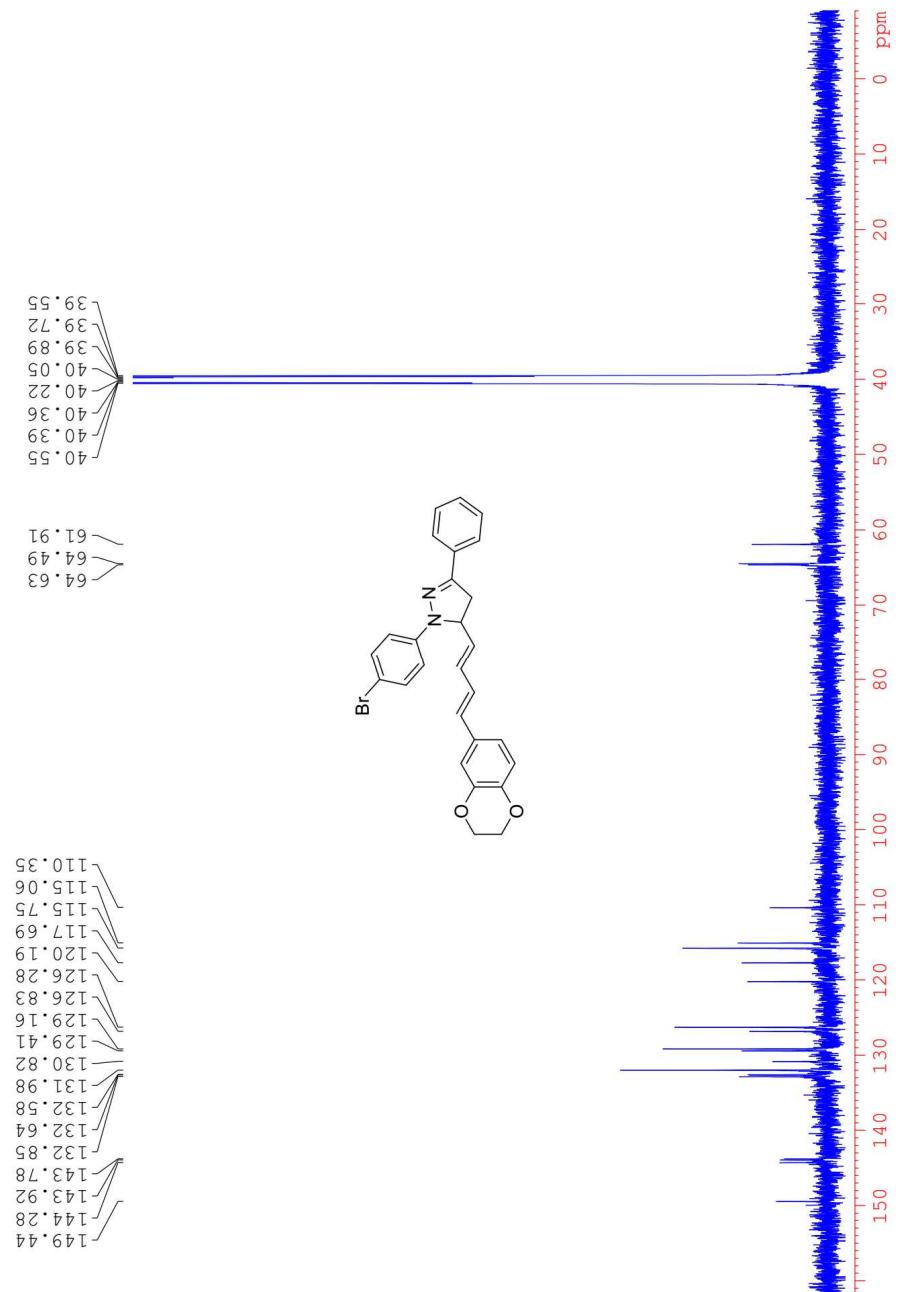
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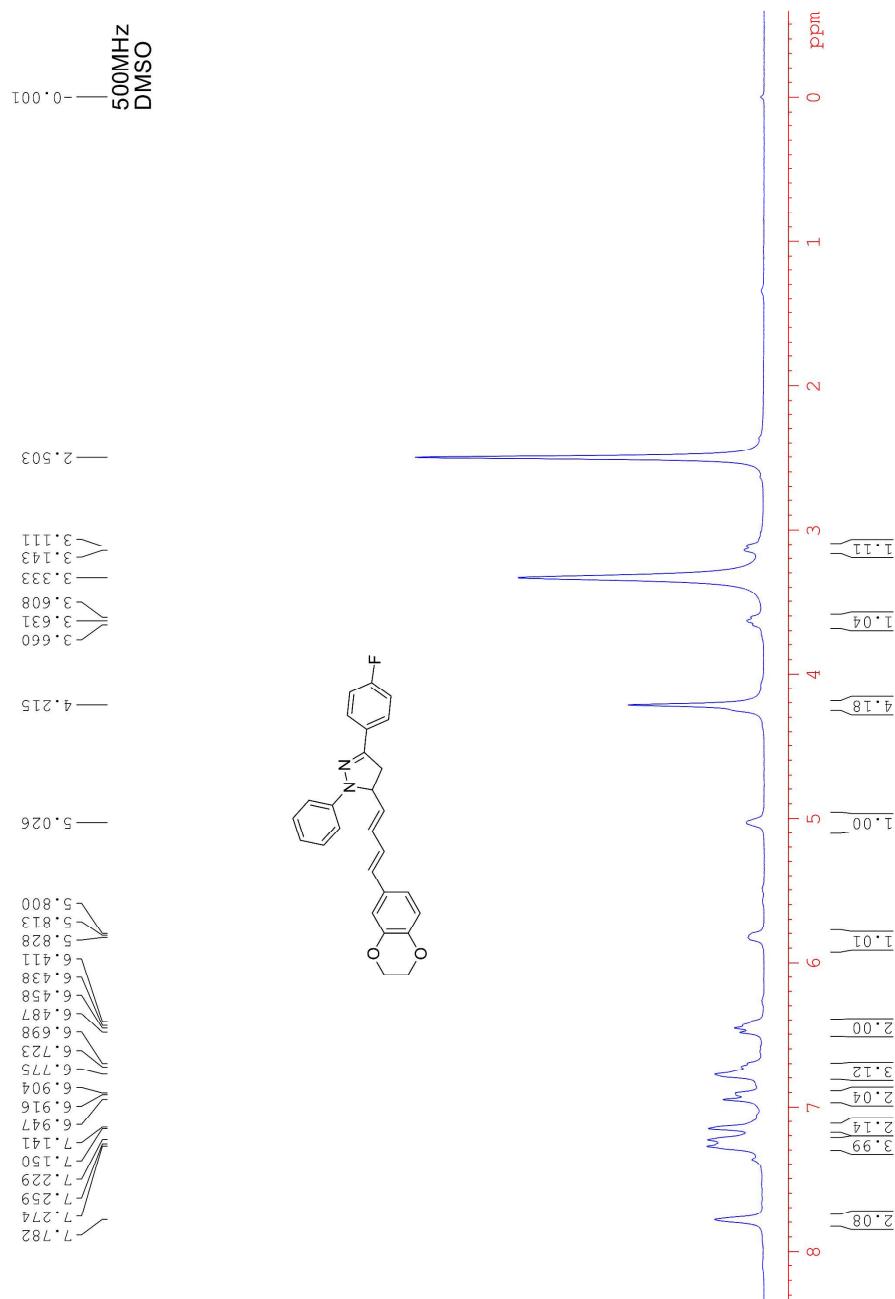
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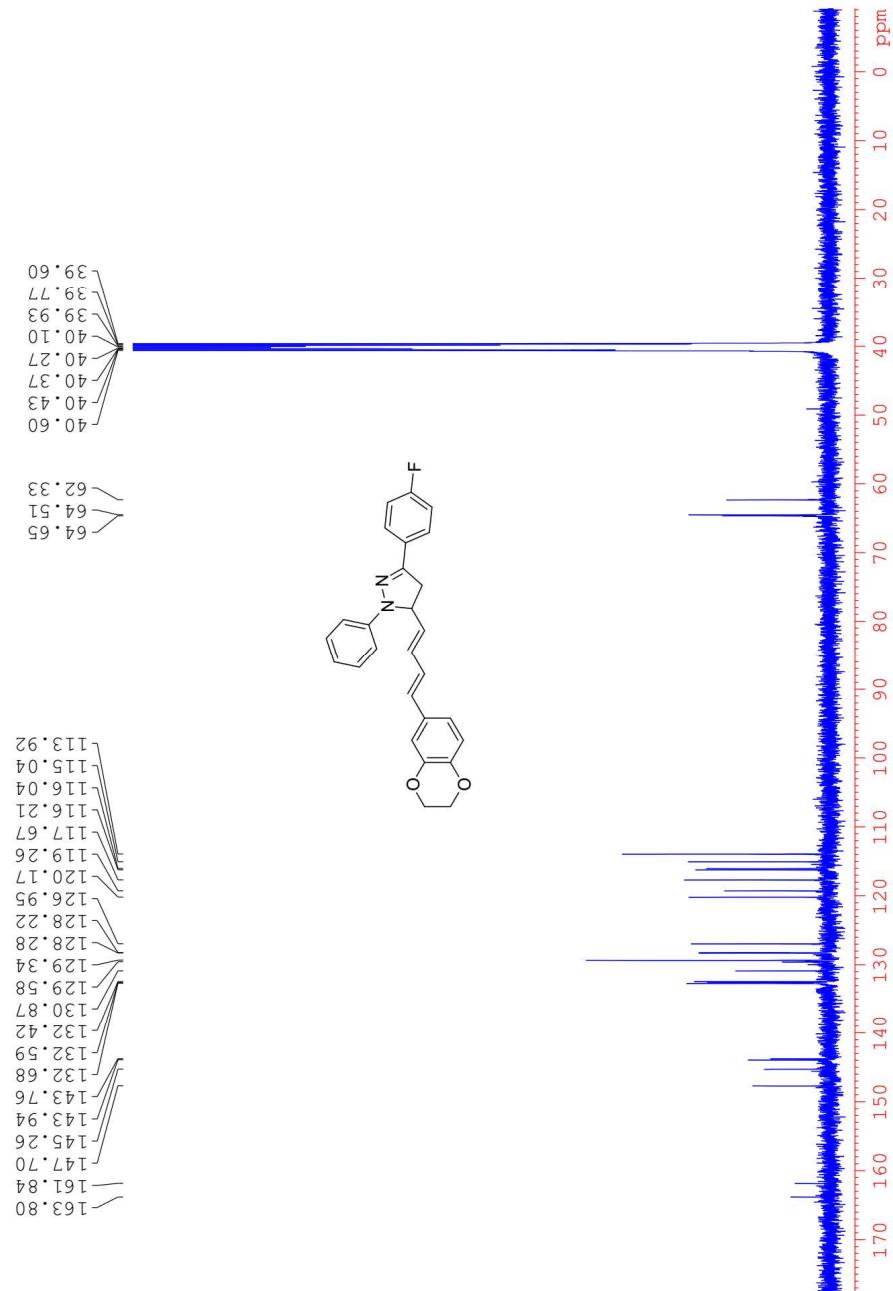
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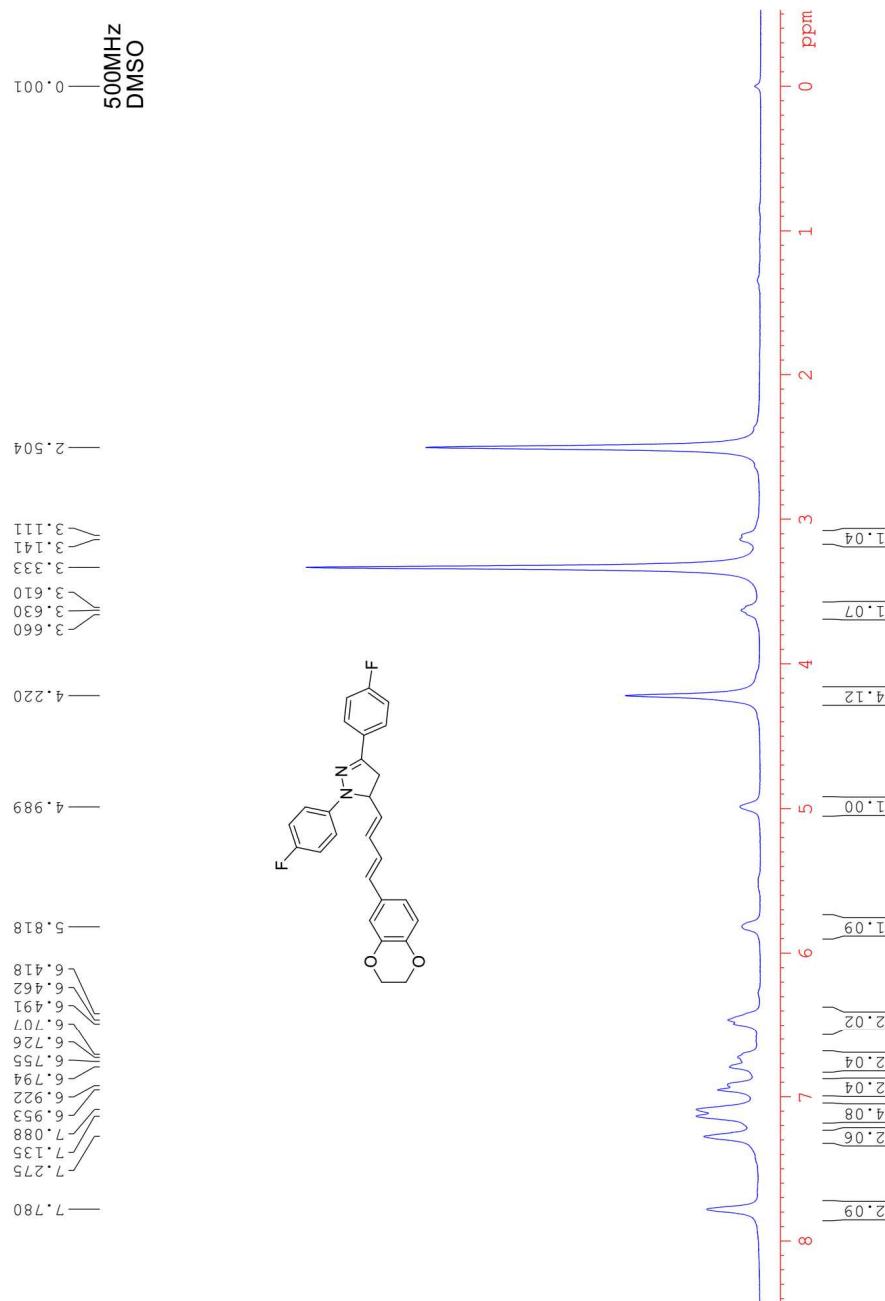
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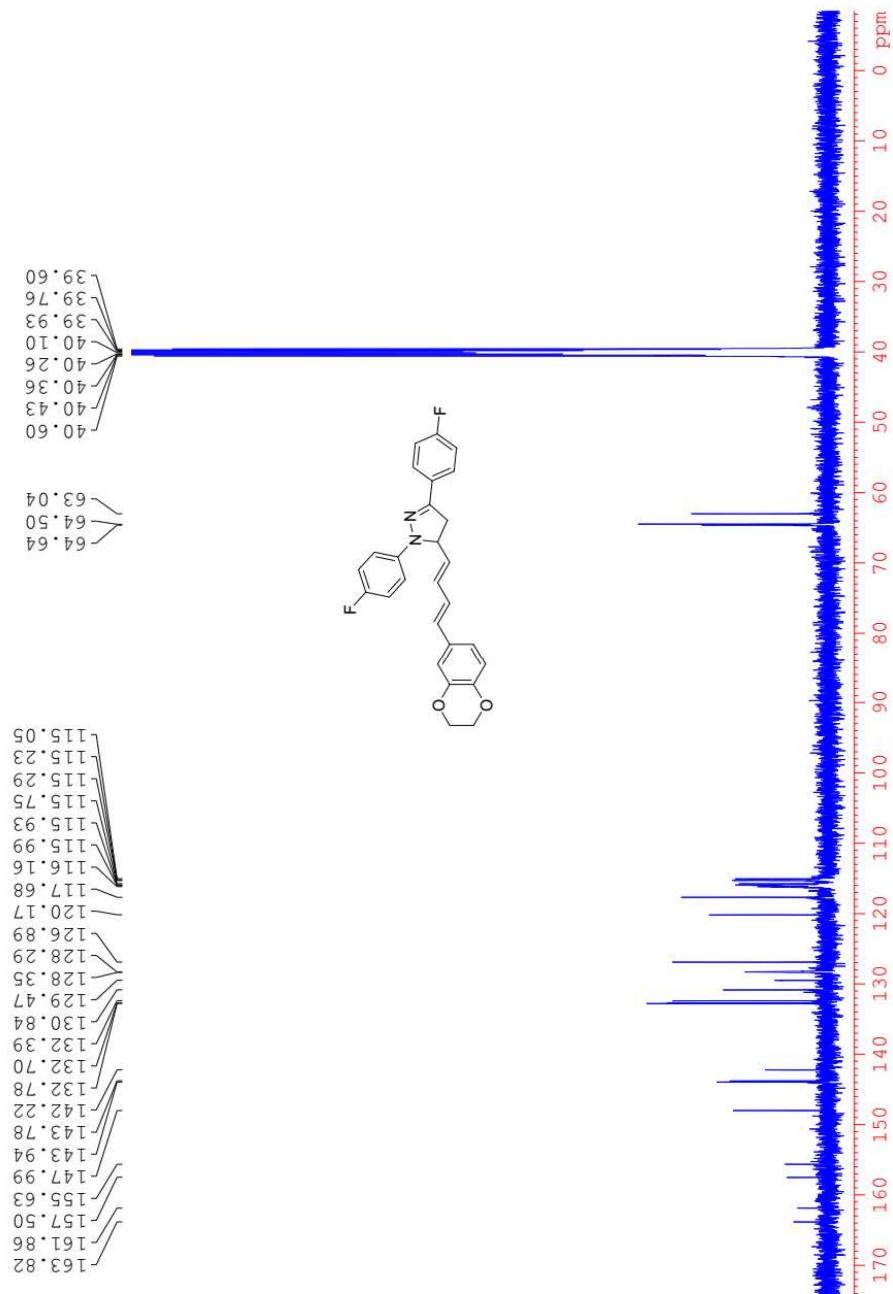
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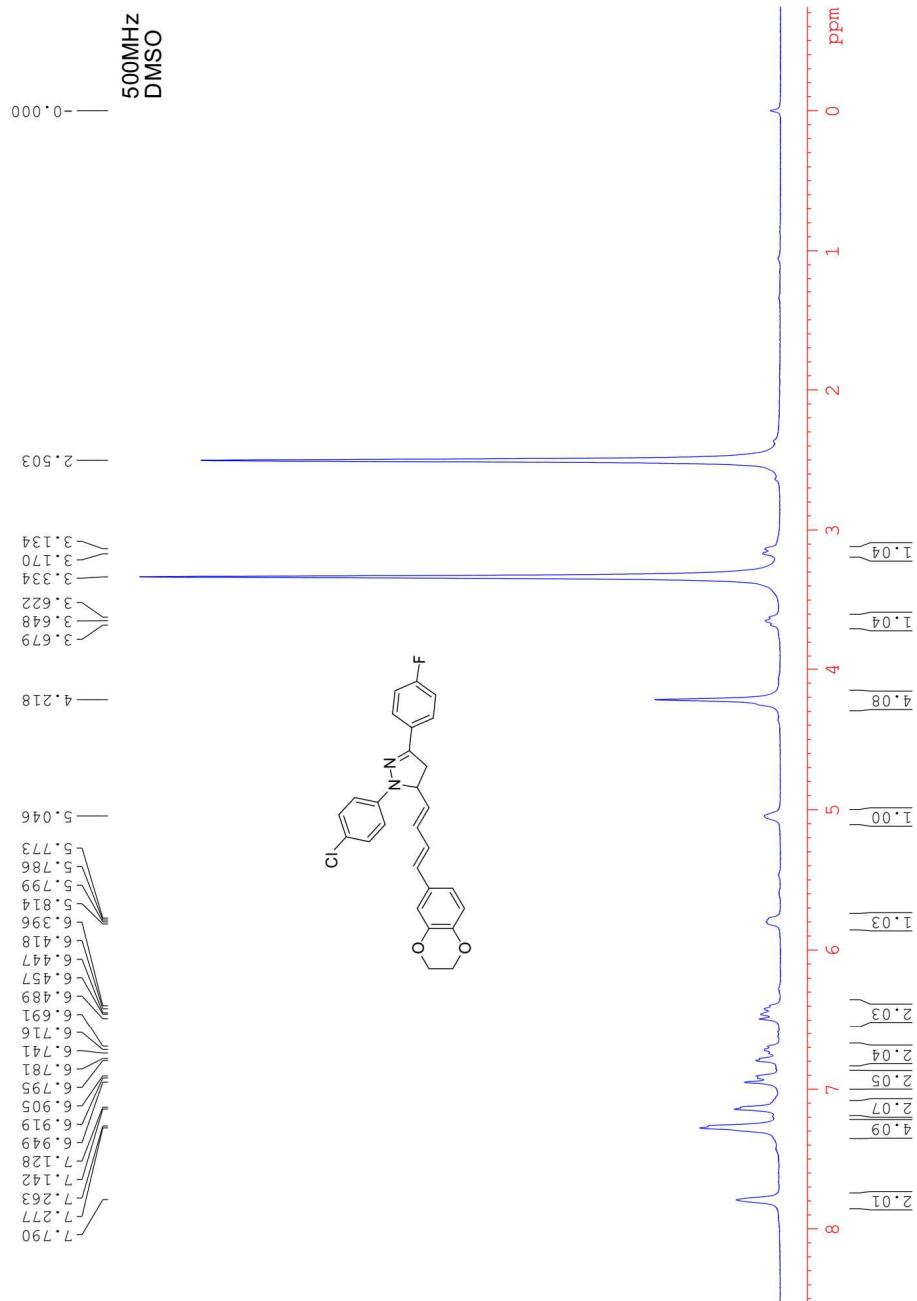
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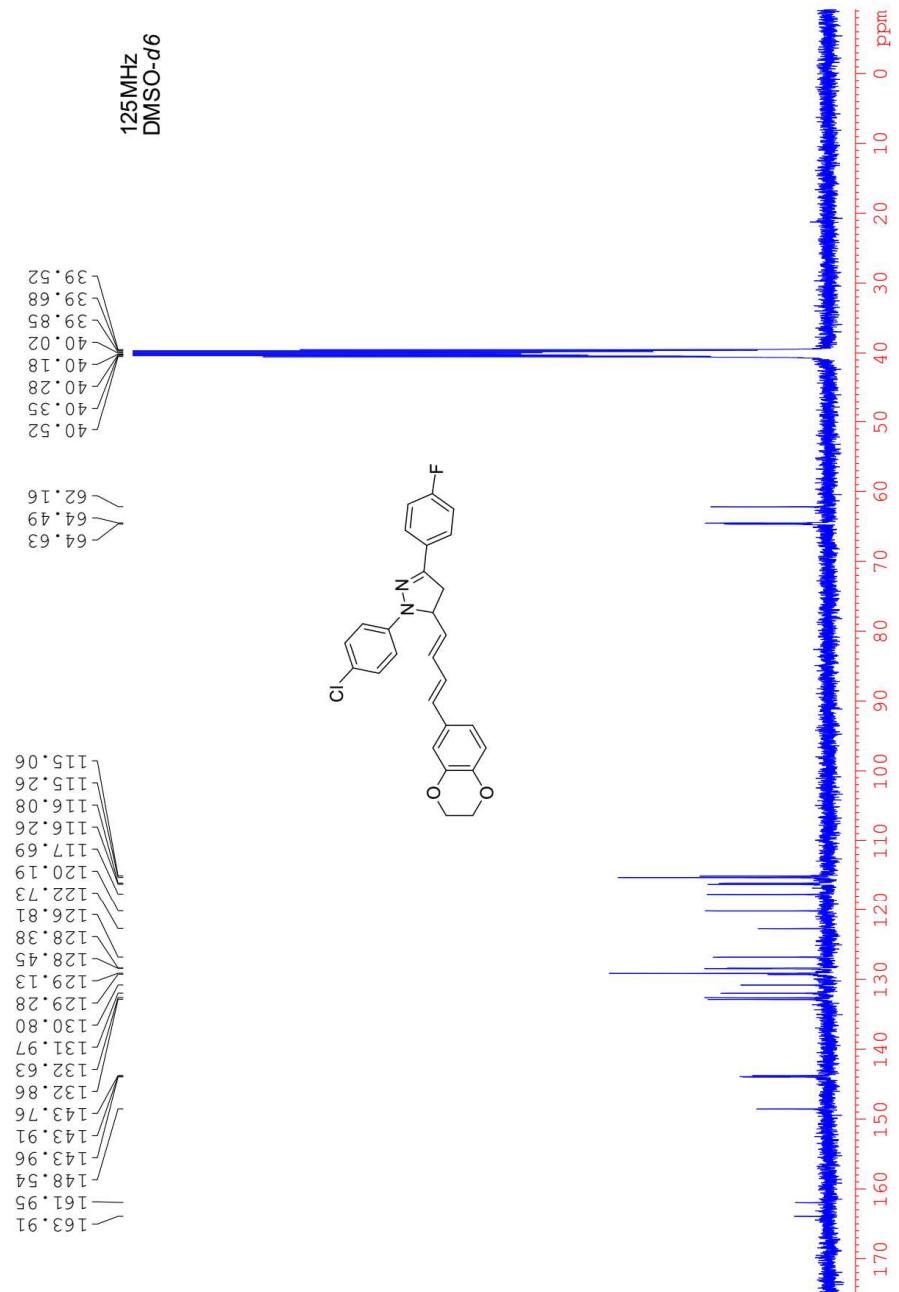
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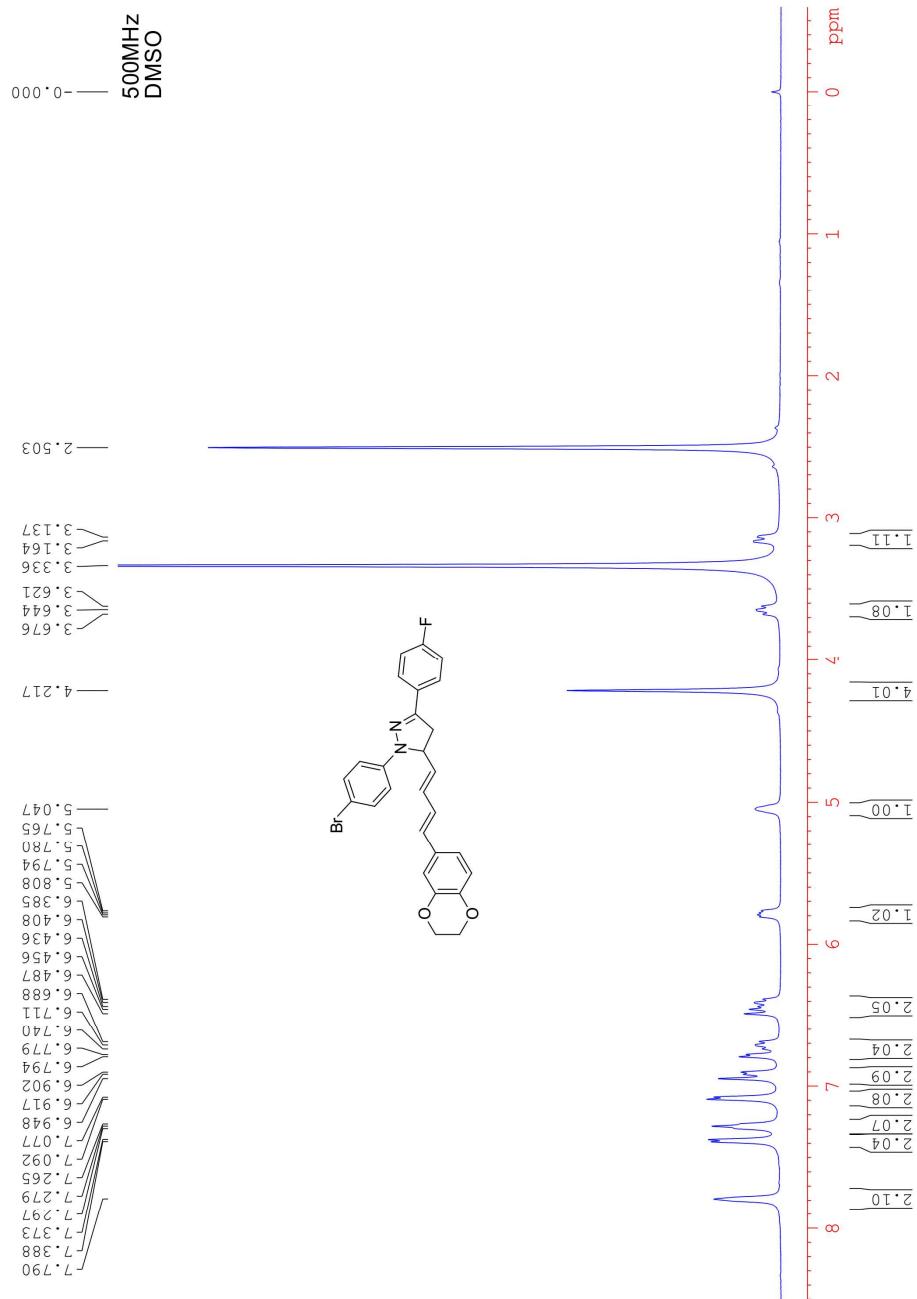
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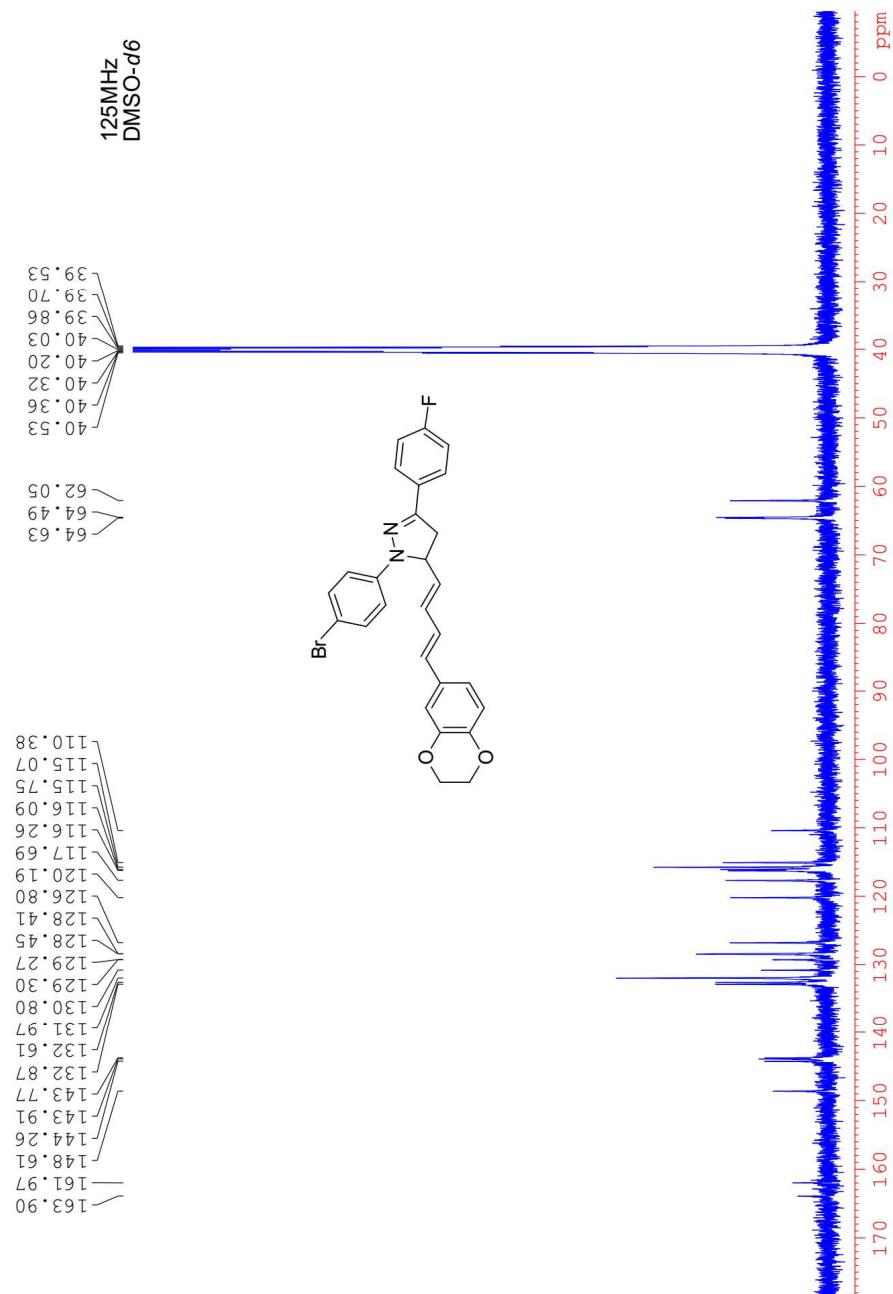
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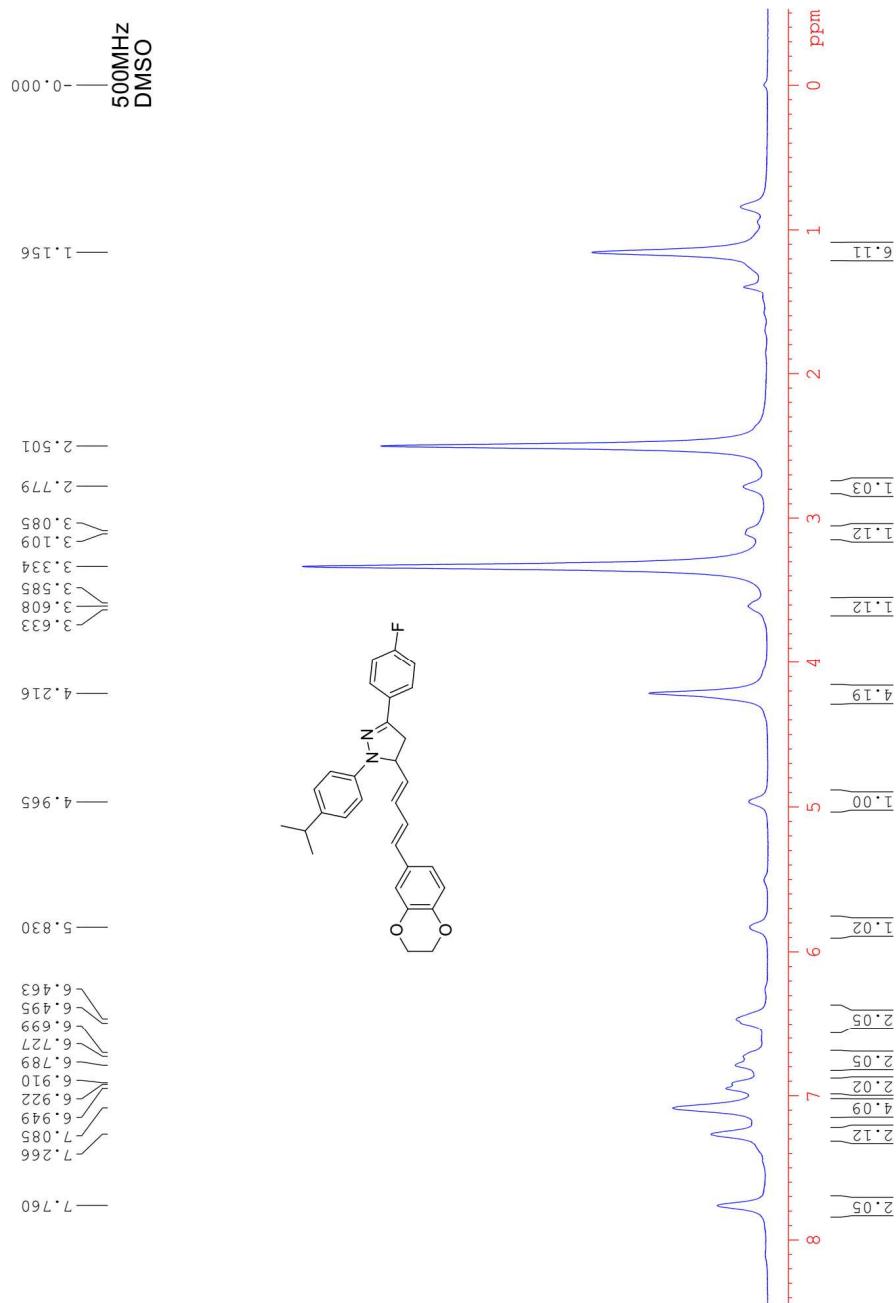
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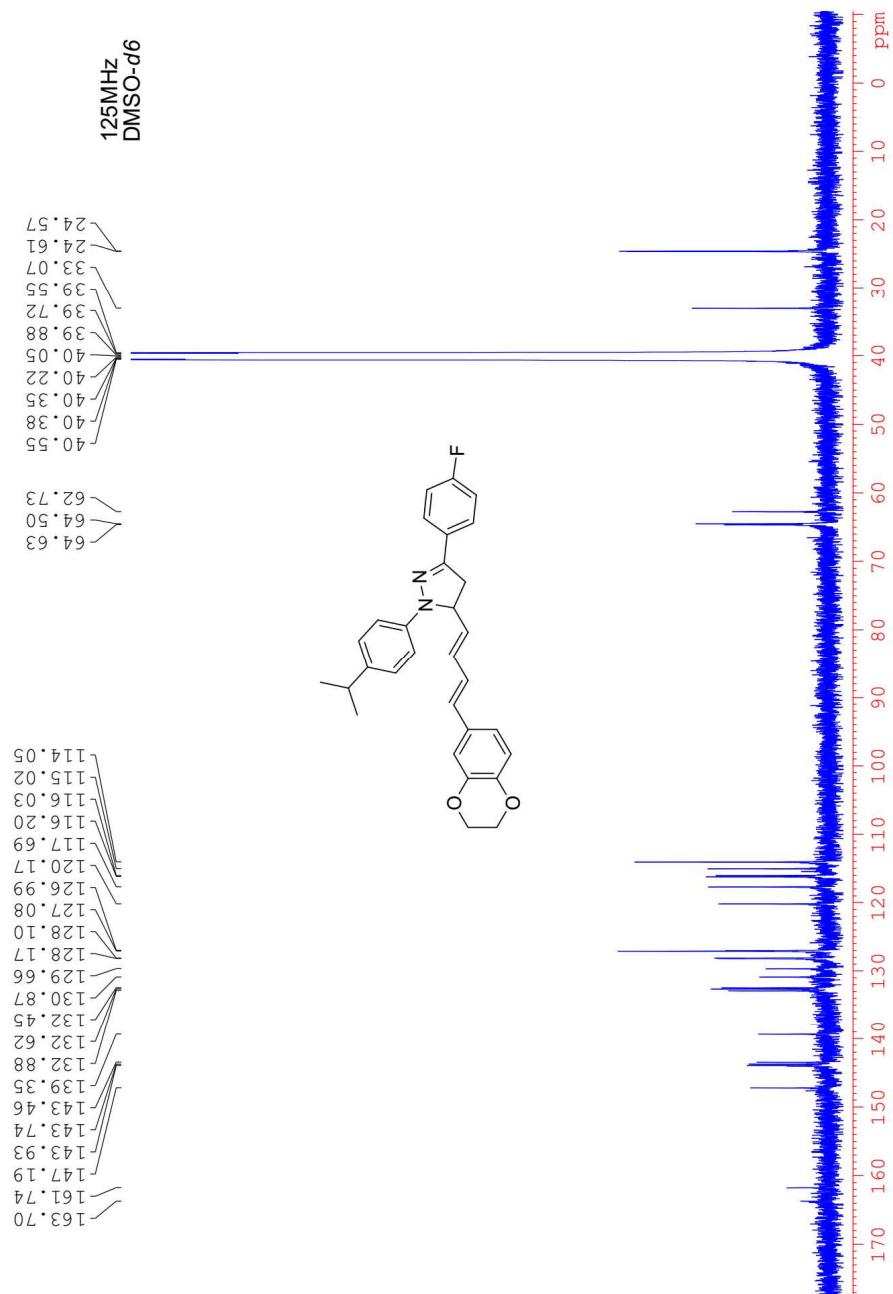
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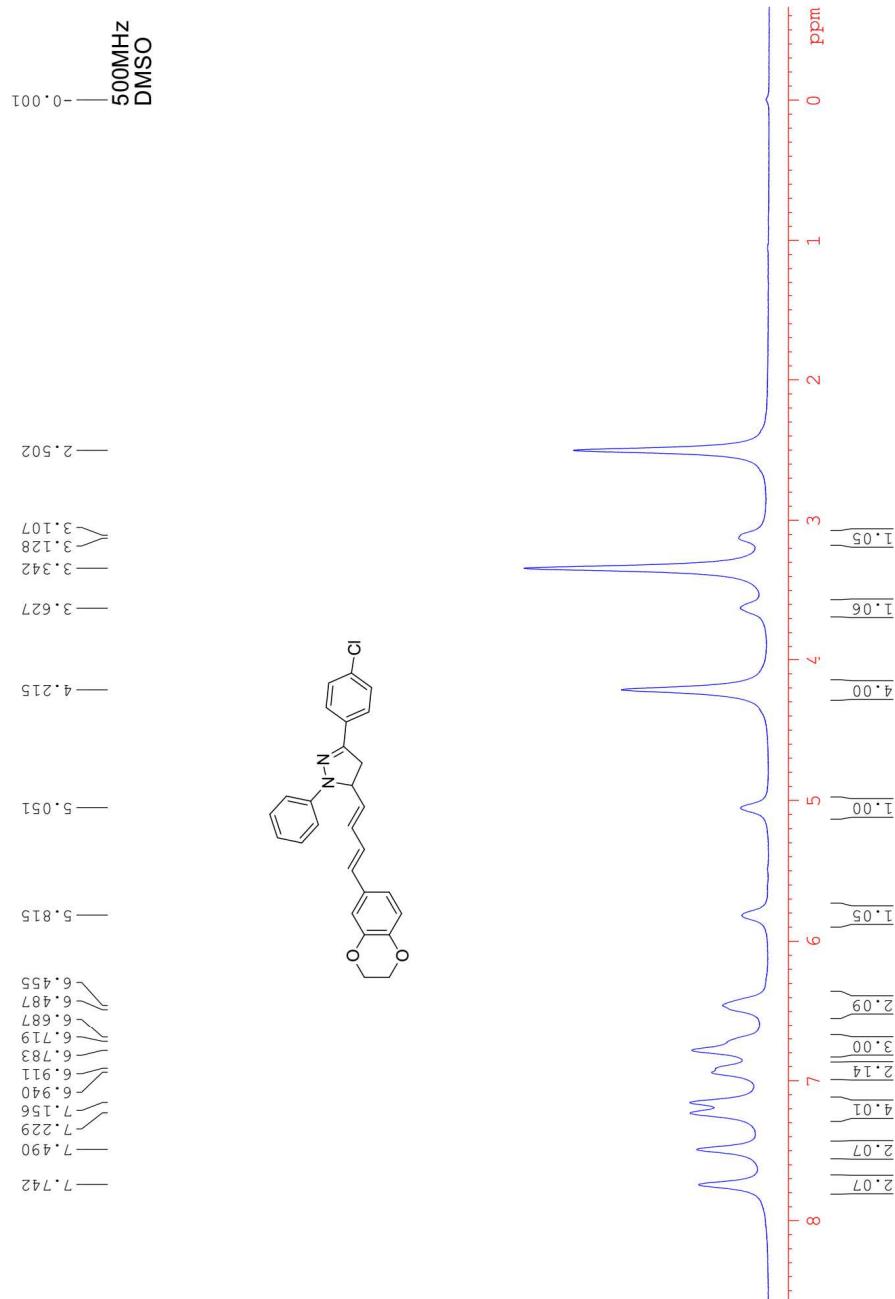
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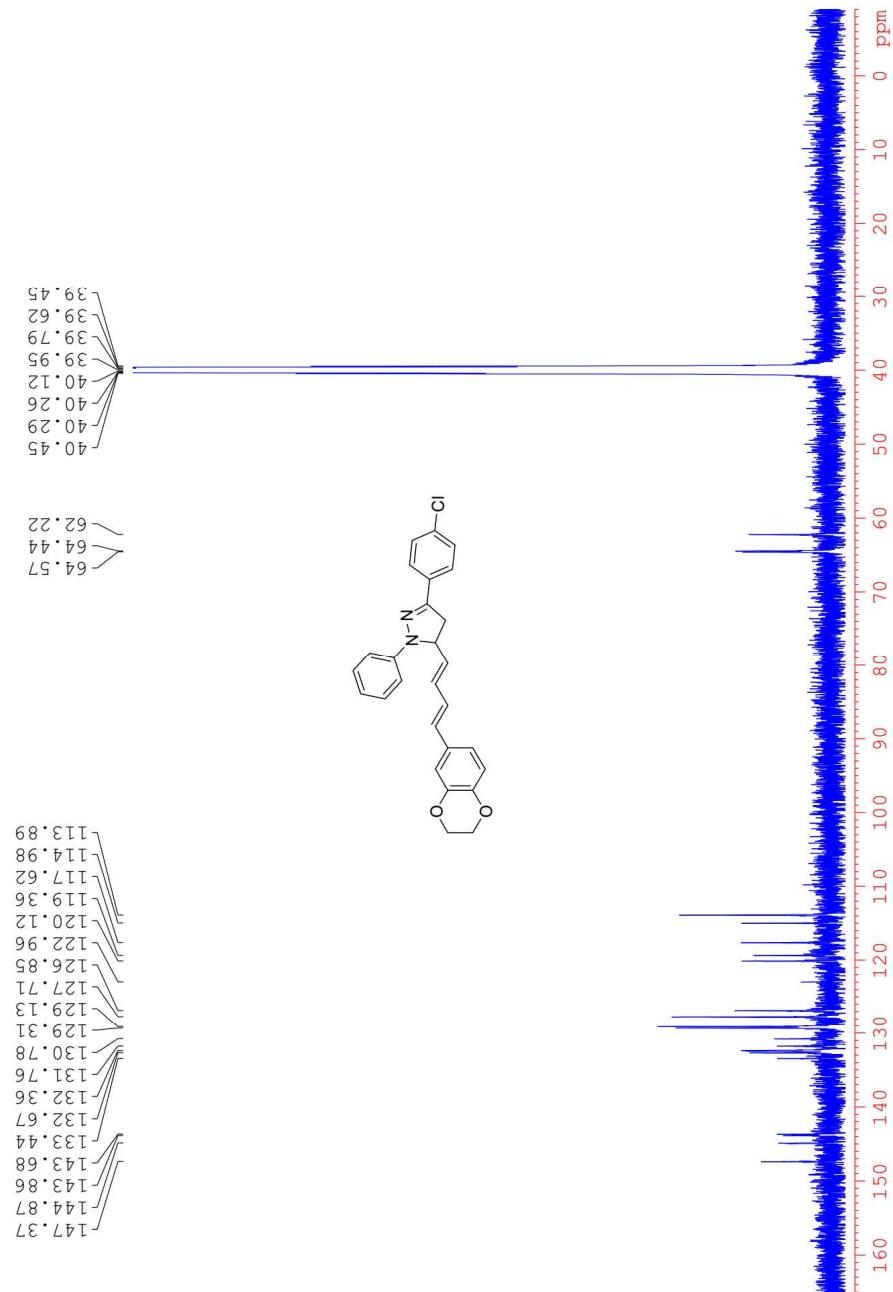
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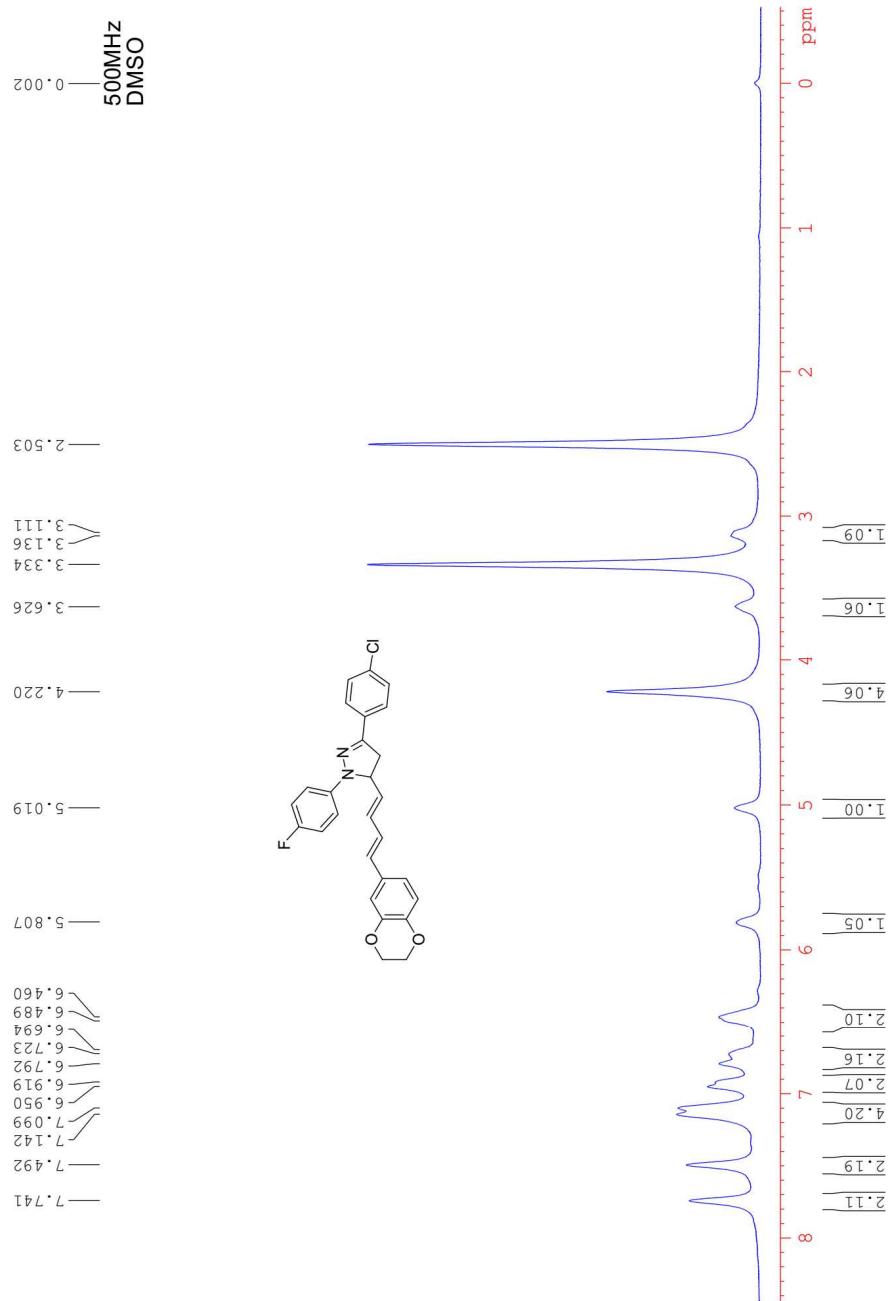
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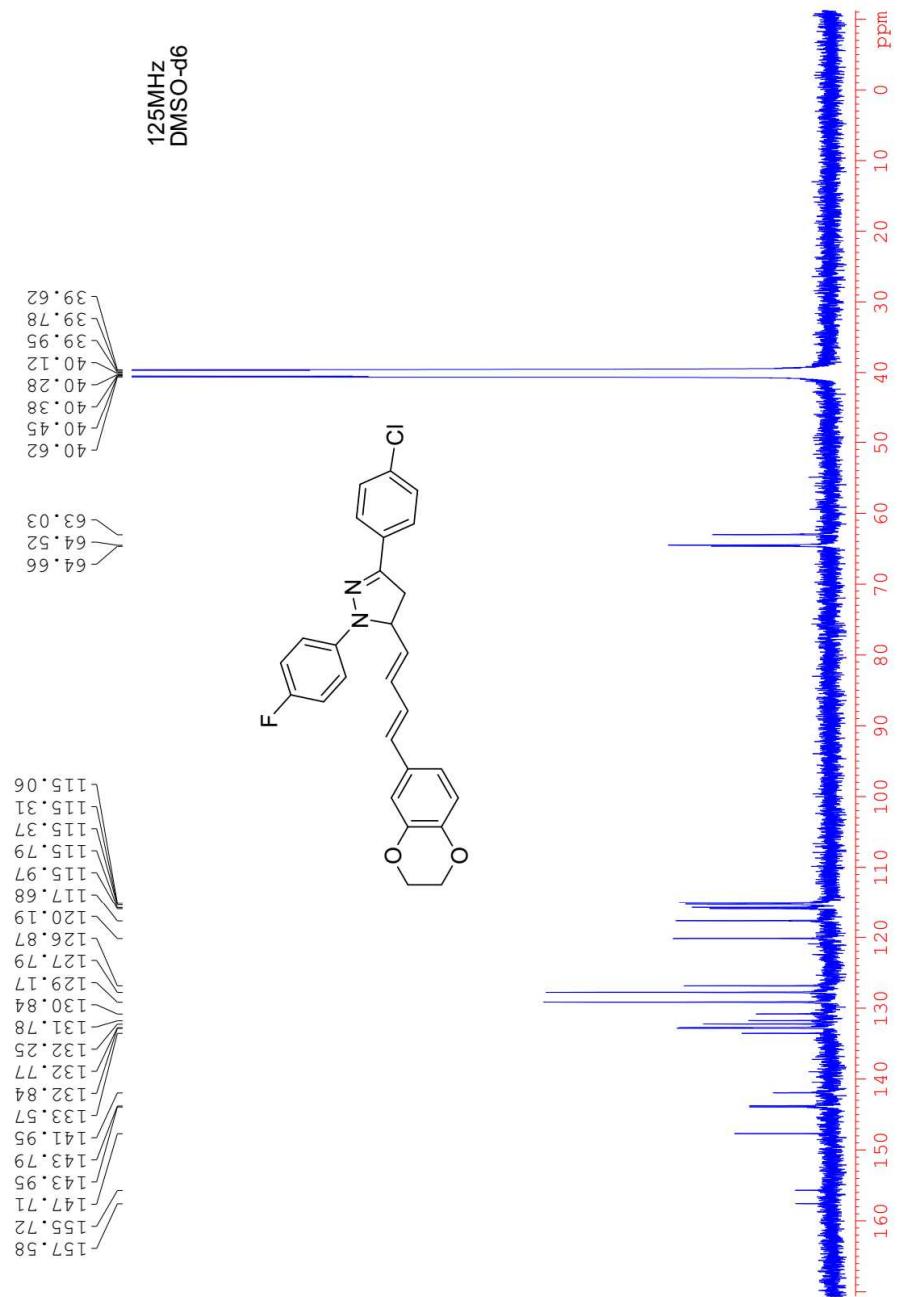
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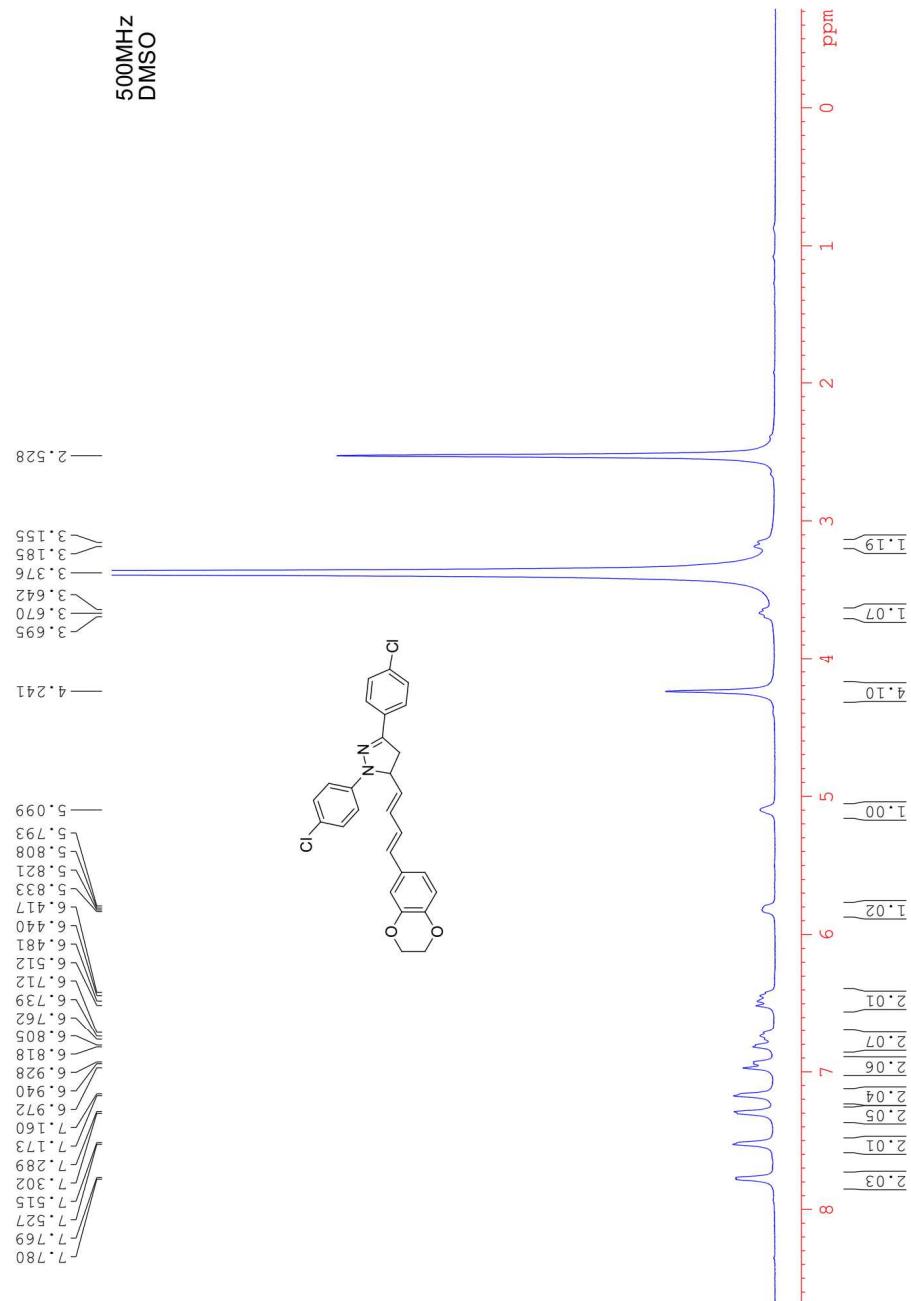
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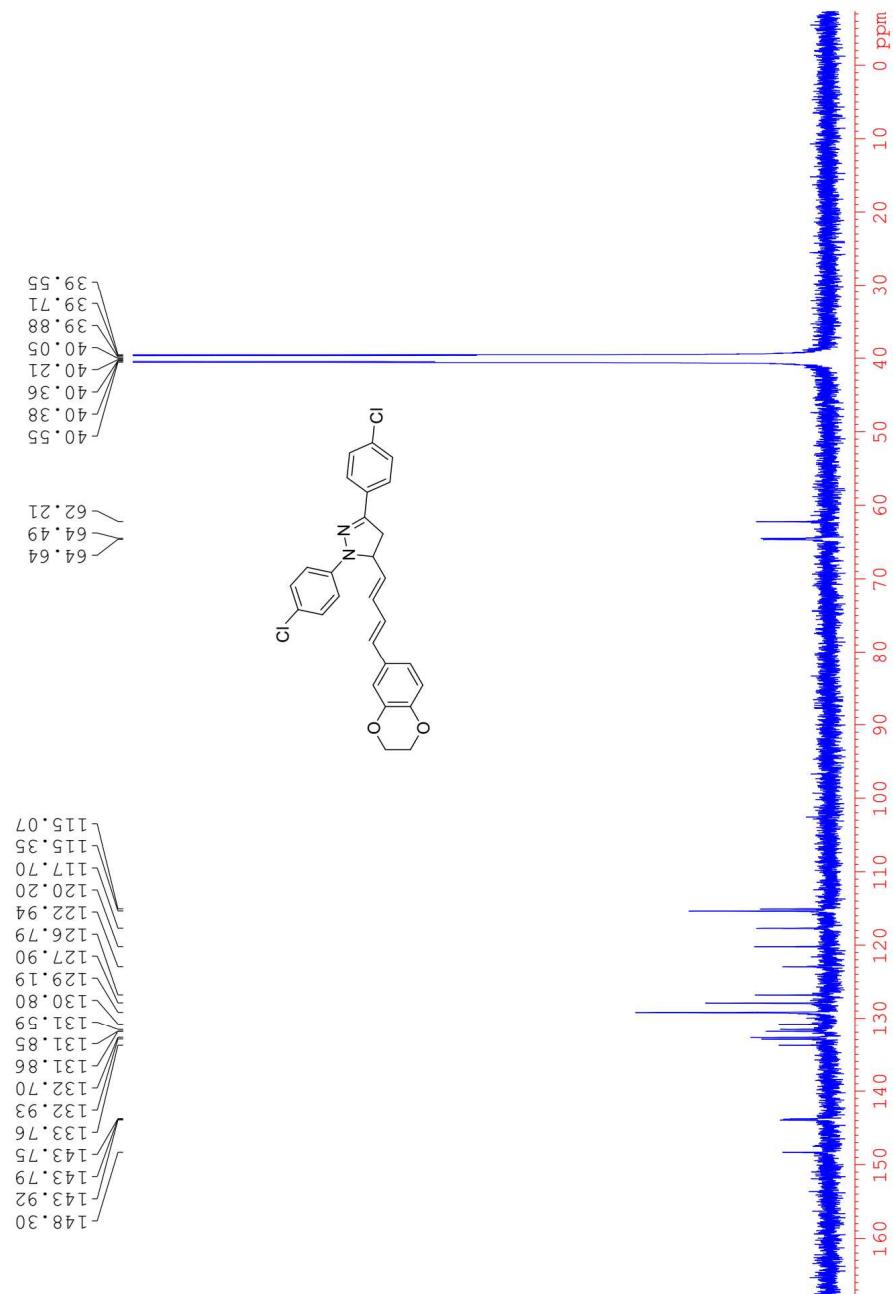
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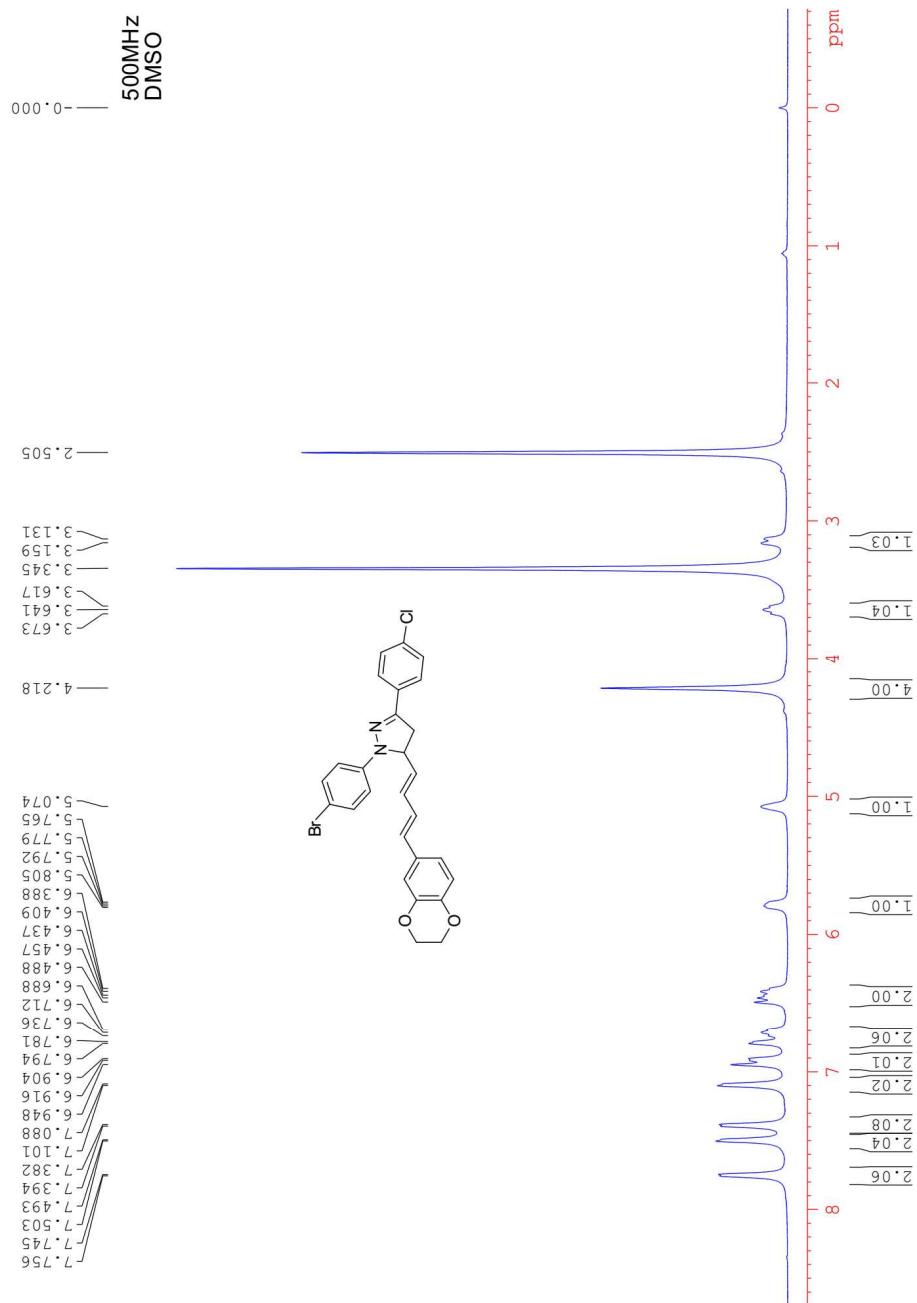
compound IXl



compound IXl



compound IXm



compound IXm

