

**Enantioselective Cascade Reaction between α,β -Unsaturated Carbonyls and
Malonic Half-thioesters: Rapid Access to Chiral δ -lactones**

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Supporting Information

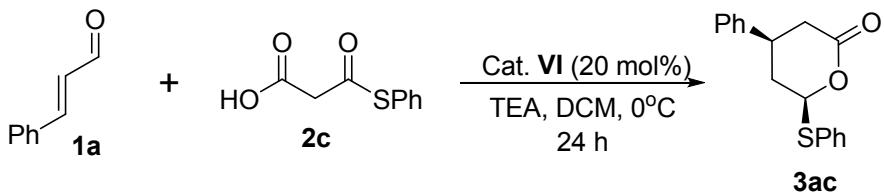
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1. General Information

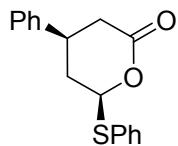
Chemicals and solvents were purchased from commercial suppliers and used as received. ^1H and ^{13}C NMR spectra were recorded on a Bruker ACF300 (300 MHz) or AMX500 (500 MHz) spectrometer. Chemical shifts were reported in parts per million (ppm), and the residual solvent peak was used as an internal reference: proton (chloroform δ 7.26), carbon (chloroform δ 77.0) or tetramethylsilane (TMS δ 0.00) was used as a reference. Multiplicity was indicated as follows: s (singlet), d (doublet), t (triplet), q (quartet), m (multiplet), dd (doublet of doublet), bs (broad singlet). Coupling constants were reported in Hertz (Hz). Low resolution mass spectra were obtained on a Finnigan/MAT LCQ spectrometer in ESI mode, and a Finnigan/MAT 95XL-T mass spectrometer in EI mode. All high resolution mass spectra were obtained on a Finnigan/MAT 95XL-T spectrometer. For thin layer chromatography (TLC), Merck pre-coated TLC plates (Merck 60 F254) were used, and compounds were visualized with a UV light at 254 nm. Flash chromatography separations were performed on Merck 60 (0.040-0.063 mm) mesh silica gel. The enantiomeric excesses of products were determined by chiral phase HPLC analysis. Optical rotations were recorded on Jasco DIP-1000 polarimeter.

2. Experimental Procedure for Decarboxylative Reaction of MAHTs to Enals

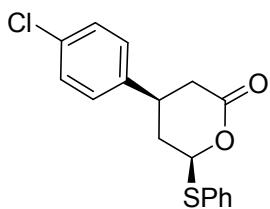


Typical procedure for the decarboxylative reaction:

To a solution of cinnamaldehyde **1a** (26.4 mg, 25.2 uL, 0.2 mmol,) in DCM (0.5 mL) was added catalyst **V** (25.6 mg, 0.04 mmol) at 0 °C. After 40 min, malonic acid half thioester **2c** (98.1 mg, 0.5 mmol) was added in one portion. Then, TEA (50.5 mg, 69.6 uL, 0.5 mmol) in 0.5 mL DCM was added dropwise *via* syringe in 10 min at 0 °C. The resulting reaction mixture was kept stirring at 0°C for 24 h. The crude product was purified by silica gel flash chromatography, eluted by hexane/EtOAc = 10:1 to afford the desired product **3ac** as white solid (46.7 mg, 82% yield).

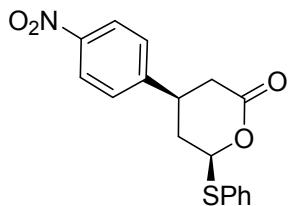


(4S,6S)-4-phenyl-6-(phenylthio)tetrahydro-2H-pyran-2-one (3ac) (Table 2, entry 1). 46.7 mg, 82% yield; White solid; ¹H NMR (500 MHz, CDCl₃) δ (ppm): 7.61 – 7.59 (m, 2H), 7.36 (ddd, *J* = 7.7, 4.0, 1.7 Hz, 5H), 7.30 – 7.26 (m, 1H), 7.17 – 7.16 (m, 2H), 5.74 (dd, *J* = 11.3, 4.1 Hz, 1H), 3.19 (tt, *J* = 12.3, 4.4 Hz, 1H), 2.88 (ddd, *J* = 17.7, 5.1, 2.3 Hz, 1H), 2.58 – 2.51 (m, 2H), 2.04 – 1.97 (m, 1H); ¹³C NMR (125 MHz, CDCl₃) δ (ppm): 169.4, 141.5, 133.3, 131.8, 129.2, 129.1, 128.6, 127.5, 126.4, 86.3, 37.4, 37.2, 36.3; HRMS (EI) calcd for C₁₇H₁₆O₂S 284.0871, found 284.0876; HPLC (Chiraldak IC, *i*-propanol/hexane = 20/80, flow rate 1.0 mL/min, λ = 254 nm): *t*_R (major) = 28.2 min, *t*_R (minor) = 23.0 min, *ee* = 92%; [α]²⁵_D = -96.6 (*c* = 1.08 in DCM).



(4*S*,6*S*)-4-(4-chlorophenyl)-6-(phenylthio)tetrahydro-2*H*-pyran-2-one (3bc) (Table 2, entry 2).

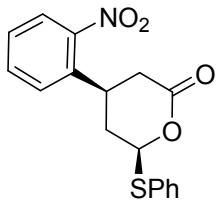
52.9 mg, 83% yield; White solid; ^1H NMR (500 MHz, CDCl_3) δ (ppm): 7.60 – 7.58 (m, 2H), 7.38 – 7.34 (m, 3H), 7.34 – 7.31 (m, 2H), 7.11 – 7.09 (m, 2H), 5.72 (dd, J = 11.3, 4.1 Hz, 1H), 3.17 (tt, J = 12.2, 4.5 Hz, 1H), 2.86 (ddd, J = 17.6, 5.1, 2.2 Hz, 1H), 2.56 – 2.46 (m, 2H), 2.00 – 1.93 (m, 1H); ^{13}C NMR (125 MHz, CDCl_3) δ (ppm): 169.1, 134.0, 133.3, 133.3, 131.6, 129.2, 129.2, 128.7, 127.8, 86.2, 37.1, 36.9, 36.1; HRMS (EI) calcd for $\text{C}_{17}\text{H}_{15}\text{O}_2\text{ClS}$ 318.0481, found 318.0486; HPLC (Chiralpak IC, *i*-propanol/hexane = 20/80, flow rate 1.0 mL/min, λ = 254 nm): t_{R} (major) = 28.4 min, t_{R} (minor) = 24.7 min, ee = 94%; $[\alpha]^{25}_{\text{D}} = -52.7$ (c = 1.05 in DCM).



(4*S*,6*S*)-4-(4-nitrophenyl)-6-(phenylthio)tetrahydro-2*H*-pyran-2-one (3cc) (Table 2, entry 3). 57.3

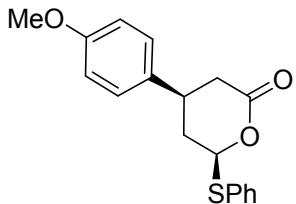
mg, 87% yield; White solid; ^1H NMR (500 MHz, CDCl_3) δ (ppm): 8.21 (d, J = 8.7 Hz, 2H), 7.60 – 7.58 (m, 2H), 7.37 – 7.34 (m, 5H), 5.74 (dd, J = 11.1, 4.1 Hz, 1H), 3.35 (tt, J = 12.1, 4.5 Hz, 1H), 2.90 (ddd, J = 17.5, 5.2, 2.2 Hz, 1H), 2.61 – 2.51 (m, 2H), 2.02 (dt, J = 13.8, 11.7 Hz, 1H); ^{13}C NMR (125 MHz, CDCl_3) δ (ppm): 168.5, 148.7, 147.3, 133.4, 131.3, 129.2, 128.8, 127.5, 124.3, 86.0, 37.3, 36.6, 35.7; HRMS (EI) calcd for $\text{C}_{17}\text{H}_{15}\text{O}_4\text{NS}$ 329.0722, found 329.0723; HPLC (Chiralpak IA, *i*-propanol/hexane = 20/80, flow rate 1.0 mL/min, λ = 254 nm): t_{R} (major) = 36.5 min, t_{R} (minor) = 26.9 min, ee = 95%;

$[\alpha]^{25}_{\text{D}} = -68.8$ ($c = 0.97$ in DCM).



(4*S*,6*S*)-4-(2-nitrophenyl)-6-(phenylthio)tetrahydro-2*H*-pyran-2-one (3dc) (Table 2, entry 4). 52.0

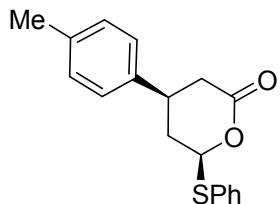
mg, 79% yield; Colorless oil; ^1H NMR (500 MHz, CDCl_3) δ (ppm): 7.87 (d, $J = 8.1$ Hz, 1H), 7.64 (t, $J = 7.6$ Hz, 1H), 7.59 (dd, $J = 6.4, 2.8$ Hz, 2H), 7.44 (t, $J = 7.7$ Hz, 1H), 7.37 – 7.35 (m, 4H), 5.74 (dd, $J = 11.3, 3.8$ Hz, 1H), 3.79 (td, $J = 11.6, 5.8$ Hz, 1H), 2.99 (ddd, $J = 17.5, 5.3, 1.8$ Hz, 1H), 2.64 – 2.54 (m, 2H), 2.03 (dd, $J = 25.3, 11.7$ Hz, 1H); ^{13}C NMR (125 MHz, CDCl_3) δ (ppm): 168.7, 149.3, 136.0, 133.7, 133.3, 131.4, 129.2, 128.7, 128.3, 127.4, 124.9, 86.0, 36.5, 35.9, 32.2; HRMS (EI) calcd for $\text{C}_{17}\text{H}_{15}\text{O}_4\text{NS}$ 329.0722, found 329.0734; HPLC (Chiralpak IA, *i*-propanol/hexane = 10/90, flow rate 1.0 mL/min, $\lambda = 254$ nm): t_{R} (major) = 30.4 min, t_{R} (minor) = 34.1 min, $ee = 96\%$; $[\alpha]^{25}_{\text{D}} = -57.2$ ($c = 1.00$ in DCM).



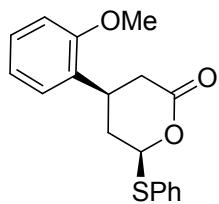
(4*S*,6*S*)-4-(4-methoxyphenyl)-6-(phenylthio)tetrahydro-2*H*-pyran-2-one (3ec) (Table 2, entry 5).

51.7 mg, 82% yield; White solid; ^1H NMR (500 MHz, CDCl_3) δ (ppm): 7.59 (dd, $J = 6.4, 3.2$ Hz, 2H), 7.35 (dd, $J = 4.9, 1.9$ Hz, 3H), 7.10 – 7.04 (m, 2H), 6.90 – 6.87 (m, 2H), 5.72 (dd, $J = 11.3, 4.1$ Hz, 1H), 3.80 (s, 3H), 3.13 (tt, $J = 12.6, 3.8$ Hz, 1H), 2.86 (ddd, $J = 17.7, 5.1, 2.3$ Hz, 1H), 2.55 – 2.46 (m, 2H), 2.00 – 1.92 (m, 1H); ^{13}C NMR (125 MHz, CDCl_3) δ (ppm): 169.6, 158.8, 133.6, 133.2, 131.80, 129.2,

128.6, 127.4, 114.4, 86.4, 55.3, 37.5, 36.7, 36.5; HRMS (EI) calcd for $C_{18}H_{18}O_3S$ 314.0977, found 314.0987; HPLC (Chiralpak IC, *i*-propanol/hexane = 20/80, flow rate 1.0 mL/min, λ = 254 nm): t_R (major) = 42.9 min, t_R (minor) = 38.0 min, ee = 88%; $[\alpha]^{25}_D$ = -74.7 (c = 0.91 in DCM).

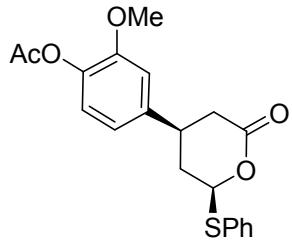


(4S,6S)-6-(phenylthio)-4-p-tolyltetrahydro-2H-pyran-2-one (3fc) (Table 2, entry 6). 48.6 mg, 81% yield; Colorless oil; 1H NMR (500 MHz, $CDCl_3$) δ (ppm): 7.60 (dd, J = 6.4, 3.1 Hz, 2H), 7.38 – 7.33 (m, 3H), 7.16 (d, J = 8.0 Hz, 2H), 7.05 (d, J = 8.0 Hz, 2H), 5.73 (dd, J = 11.3, 4.1 Hz, 1H), 3.15 (tt, J = 12.0, 4.4 Hz, 1H), 2.86 (ddd, J = 17.7, 5.1, 2.3 Hz, 1H), 2.55 – 2.48 (m, 2H), 2.34 (s, 3H), 2.02 – 1.94 (m, 1H); ^{13}C NMR (125 MHz, $CDCl_3$) δ (ppm): 169.6, 138.6, 137.2, 133.2, 131.8, 129.7, 129.1, 128.6, 126.2, 86.5, 37.3, 37.1, 36.4, 21.0; HRMS (EI) calcd for $C_{18}H_{18}O_2S$ 298.1028, found 298.1028; HPLC (Chiralpak IC, *i*-propanol/hexane = 20/80, flow rate 1.0 mL/min, λ = 254 nm): t_R (major) = 25.8 min, t_R (minor) = 22.5 min, ee = 93%; $[\alpha]^{25}_D$ = -35.2 (c = 1.05 in DCM).

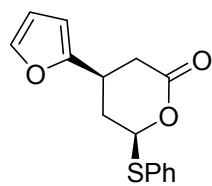


(4S,6S)-4-(2-methoxyphenyl)-6-(phenylthio)tetrahydro-2H-pyran-2-one (3gc) (Table 2, entry 7). 58.3 mg, 93% yield; Colorless oil; 1H NMR (500 MHz, $CDCl_3$) δ (ppm): 7.63 – 7.59 (m, 2H), 7.36 – 7.33 (m, 3H), 7.26 (td, J = 8.0, 1.7 Hz, 1H), 7.06 (dd, J = 7.5, 1.6 Hz, 1H), 6.94 (td, J = 7.5, 0.9 Hz, 1H), 6.88 (d, J = 8.2 Hz, 1H), 5.74 (dd, J = 11.4, 3.9 Hz, 1H), 3.81 (s, 3H), 3.53 (tt, J = 12.0, 4.7, 1H),

2.87 (ddd, $J = 17.6, 5.4, 2.1$ Hz, 1H), 2.57 (dd, $J = 17.6, 11.7$ Hz, 1H), 2.49 (dtd, $J = 13.8, 3.9, 2.1$ Hz, 1H), 2.11 – 2.03 (m, 1H); ^{13}C NMR (125 MHz, CDCl_3) δ (ppm): 170.2, 156.8, 133.1, 132.1, 129.6, 129.1, 128.4, 128.4, 126.5, 120.8, 110.7, 86.4, 55.2, 35.5, 34.5, 32.0; HRMS (EI) calcd for $\text{C}_{18}\text{H}_{18}\text{O}_3\text{S}$ 314.0977, found 314.0977; HPLC (Chiralpak IC, *i*-propanol/hexane = 20/80, flow rate 1.0 mL/min, λ = 254 nm): t_{R} (major) = 20.6 min, t_{R} (minor) = 24.6 min, $ee = 94\%$; $[\alpha]^{25}_{\text{D}} = -25.3$ ($c = 1.37$ in DCM).

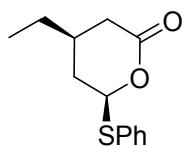


2-methoxy-4-((4S,6S)-2-oxo-6-(phenylthio)tetrahydro-2H-pyran-4-yl)phenyl acetate (3hc) (Table 2, entry 8). 60.8 mg, 82% yield; White solid; ^1H NMR (500 MHz, CDCl_3) δ (ppm): 7.61 – 7.58 (m, 2H), 7.37 – 7.34 (m, 3H), 7.02 – 6.99 (m, 1H), 6.73 (dd, $J = 6.7, 1.9$ Hz, 2H), 5.72 (dd, $J = 11.3, 4.1$ Hz, 1H), 3.82 (s, 3H), 3.18 (tt, $J = 12.2, 4.4$ Hz, 1H), 2.88 (ddd, $J = 17.6, 5.1, 2.2$ Hz, 1H), 2.60 – 2.47 (m, 2H), 2.31 (s, 3H), 2.04 – 1.91 (m, 1H); ^{13}C NMR (125 MHz, CDCl_3) δ (ppm): 169.3, 169.0, 151.4, 140.4, 138.9, 133.3, 131.6, 129.2, 128.7, 123.2, 118.3, 110.7, 86.2, 55.9, 37.3, 37.2, 36.3, 20.6; HRMS (EI) calcd for $\text{C}_{20}\text{H}_{20}\text{O}_5\text{S}$ 372.1031, found 372.1024; HPLC (Chiralpak IA, *i*-propanol/hexane = 10/90, flow rate 1.0 mL/min, λ = 254 nm): t_{R} (major) = 36.7 min, t_{R} (minor) = 40.9 min, $ee = 92\%$; $[\alpha]^{25}_{\text{D}} = -31.6$ ($c = 1.34$ in DCM).

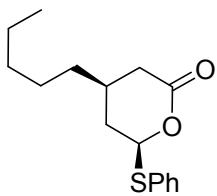


(4S,6S)-4-(furan-2-yl)-6-(phenylthio)tetrahydro-2H-pyran-2-one (3ic) (Table 2, entry 9). 43.8 mg,

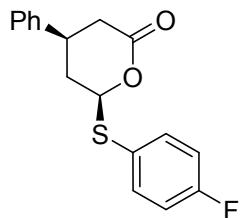
80% yield; White solid; ^1H NMR (500 MHz, CDCl_3) δ (ppm): 7.59 – 7.58 (m, 2H), 7.35 – 7.33 (m, 4H), 6.31 (dd, J = 3.2, 1.9 Hz, 1H), 6.06 (d, J = 3.2 Hz, 1H), 5.70 (dd, J = 11.3, 4.0 Hz, 1H), 3.32 (tt, J = 11.8, 4.6 Hz, 1H), 2.94 (ddd, J = 17.7, 5.3, 2.1 Hz, 1H), 2.66 (dtd, J = 14.1, 4.0, 2.2 Hz, 1H), 2.57 (dd, J = 17.7, 11.9 Hz, 1H), 1.95 (dt, J = 14.1, 11.6 Hz, 1H); ^{13}C NMR (125 MHz, CDCl_3) δ (ppm): 168.8, 154.4, 142.1, 133.4, 131.6, 129.2, 128.7, 110.3, 105.0, 86.0, 34.2, 33.9, 31.2; HRMS (EI) calcd for $\text{C}_{15}\text{H}_{14}\text{O}_3\text{S}$ 274.0664, found 274.0667; HPLC (Chiralpak IC, *i*-propanol/hexane = 20/80, flow rate 1.0 mL/min, λ = 254 nm): t_R (major) = 23.3 min, t_R (minor) = 19.0 min, ee = 91%; $[\alpha]^{25}_D$ = -67.7 (c = 0.96 in DCM).



(4*S*,6*S*)-4-ethyl-6-(phenylthio)tetrahydro-2*H*-pyran-2-one (3jc) (Table 2, entry 10). 33.4 mg, 71% yield; Colorless oil; ^1H NMR (500 MHz, CDCl_3) δ (ppm): 7.58 – 7.56 (m, 2H), 7.35 – 7.32 (m, 3H), 5.59 (dd, J = 11.2, 4.0 Hz, 1H), 2.68 (ddd, J = 17.5, 5.2, 2.0 Hz, 1H), 2.36 (dtd, J = 13.9, 4.0, 2.2 Hz, 1H), 2.05 (dd, J = 17.5, 11.6 Hz, 1H), 1.91 – 1.82 (m, 1H), 1.48 (dt, J = 13.9, 11.4 Hz, 1H), 1.37 (p, J = 7.3, 2H), 0.93 (t, J = 7.4 Hz, 3H); ^{13}C NMR (125 MHz, CDCl_3) δ (ppm): 170.1, 133.0, 132.1, 129.1, 128.4, 86.4, 35.9, 34.8, 33.3, 28.7, 10.7; HRMS (EI) calcd for $\text{C}_{15}\text{H}_{16}\text{O}_2\text{S}$ 236.0871, found 236.0870; HPLC (Chiralpak IC, *i*-propanol/hexane = 10/90, flow rate 1.0 mL/min, λ = 254 nm): t_R (major) = 35.2 min, t_R (minor) = 40.4 min, ee = 94%; $[\alpha]^{25}_D$ = -11.9 (c = 0.86 in DCM).

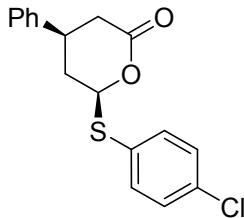


(4*S*,6*S*)-4-pentyl-6-(phenylthio)tetrahydro-2*H*-pyran-2-one (3kc) (Table 2, entry 11). 38.2 mg, 69% yield; Colorless oil; ^1H NMR (500 MHz, CDCl_3) δ (ppm): 7.57 – 7.55 (m, 2H), 7.34 – 7.32 (m, 3H), 5.59 (dd, J = 11.3, 4.0 Hz, 1H), 2.67 (ddd, J = 17.4, 5.0, 2.0 Hz, 1H), 2.37 – 2.32 (m, 2H), 2.05 (dd, J = 17.4, 11.6 Hz, 1H), 1.95 – 1.91 (m, 1H), 1.48 (dt, J = 13.8, 11.4 Hz, 1H), 1.37 – 1.26 (m, 8H), 0.88 (t, J = 7.0 Hz, 3H); ^{13}C NMR (125 MHz, CDCl_3) δ (ppm): 170.1, 133.0, 132.1, 129.1, 128.4, 86.4, 36.2, 35.8, 35.2, 31.7, 31.6, 25.9, 22.5, 14.0; HRMS (EI) calcd for $\text{C}_{16}\text{H}_{22}\text{O}_2\text{S}$ 278.1341, found 278.1341; HPLC (Chiralpak IC, *i*-propanol/hexane = 20/80, flow rate 1.0 mL/min, λ = 254 nm): t_{R} (major) = 18.1 min, t_{R} (minor) = 20.7 min, *ee* = 94%; $[\alpha]^{25}_{\text{D}} = -20.6$ (c = 0.89 in DCM).



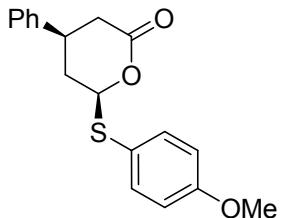
(4*S*,6*S*)-6-(4-fluorophenylthio)-4-phenyltetrahydro-2*H*-pyran-2-one (3ad) (Table 2, entry 12). 51.6 mg, 85% yield; White solid; ^1H NMR (500 MHz, CDCl_3) δ (ppm): 7.62 – 7.59 (m, 2H), 7.36 (dd, J = 10.3, 4.7 Hz, 2H), 7.30 – 7.27 (m, 1H), 7.17 – 7.15 (m, 2H), 7.08 – 7.04 (m, 2H), 5.65 (dd, J = 11.3, 4.1 Hz, 1H), 3.19 (tt, J = 12.0, 4.4 Hz, 1H), 2.88 (ddd, J = 17.7, 5.1, 2.3 Hz, 1H), 2.56 – 2.50 (m, 2H), 1.96 (ddd, J = 13.9, 12.2, 11.4 Hz, 1H); ^{13}C NMR (125 MHz, CDCl_3) δ (ppm): 169.4, 164.3, 162.3, 141.4, 136.2, 136.2, 129.1, 127.5, 126.5, 126.4, 116.4, 116.2, 86.4, 37.4, 37.2, 36.2; HRMS (EI) calcd for $\text{C}_{17}\text{H}_{15}\text{O}_2\text{FS}$ 302.0777, found 302.0777; HPLC (Chiralpak IC, *i*-propanol/hexane = 20/80, flow rate

1.0 mL/min, $\lambda = 254$ nm): t_R (major) = 22.7 min, t_R (minor) = 16.6 min, $ee = 91\%$; $[\alpha]^{25}_D = -76.2$ ($c = 1.01$ in DCM).



(4S,6S)-6-(4-chlorophenylthio)-4-phenyltetrahydro-2H-pyran-2-one (3ae) (Table 2, entry 13).

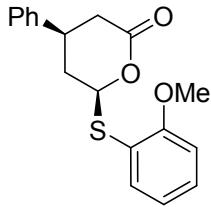
50.6 mg, 79% yield; White solid; ^1H NMR (500 MHz, CDCl_3) δ (ppm): 7.54 – 7.52 (m, 2H), 7.38 – 7.31 (m, 4H), 7.29 (d, $J = 7.5$ Hz, 1H), 7.17 (d, $J = 7.5$ Hz, 2H), 5.69 (dd, $J = 11.3, 4.1$ Hz, 1H), 3.19 (tt, $J = 12.3, 4.4$ Hz, 1H), 2.89 (ddd, $J = 17.7, 5.1, 2.2$ Hz, 1H), 2.58 – 2.52 (m, 2H), 1.98 (dt, $J = 13.8, 11.9$ Hz, 1H); ^{13}C NMR (125 MHz, CDCl_3) δ (ppm): 169.3, 141.4, 135.1, 134.7, 130.2, 129.4, 129.1, 127.6, 126.4, 86.1, 37.4, 37.2, 36.3; HRMS (EI) calcd for $\text{C}_{17}\text{H}_{15}\text{O}_2\text{ClS}$ 318.0481, found 318.0485; HPLC (Chiralpak IC, *i*-propanol/hexane = 20/80, flow rate 1.0 mL/min, $\lambda = 254$ nm): t_R (major) = 23.8 min, t_R (minor) = 18.0 min, $ee = 92\%$; $[\alpha]^{25}_D = -90.1$ ($c = 1.05$ in DCM).



(4S,6S)-6-(4-methoxyphenylthio)-4-phenyltetrahydro-2H-pyran-2-one (3af) (Table 2, entry 14).

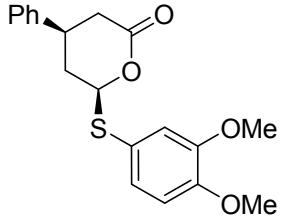
53.6 mg, 85% yield; White solid; ^1H NMR (500 MHz, CDCl_3) δ (ppm): 7.56 – 7.53 (m, 2H), 7.35 (dd, $J = 10.3, 4.7$ Hz, 2H), 7.28 (dt, $J = 3.9, 1.6$ Hz, 1H), 7.15 (d, $J = 7.2$ Hz, 2H), 6.90 – 6.88 (m, 2H), 5.59 (dd, $J = 11.2, 4.1$ Hz, 1H), 3.82 (s, 3H), 3.16 (tt, $J = 12.0, 4.5$ Hz, 1H), 2.85 (ddd, $J = 17.7, 5.1, 2.3$ Hz, 1H), 2.53 – 2.45 (m, 2H), 1.94 (ddd, $J = 13.9, 12.3, 11.4$ Hz, 1H); ^{13}C NMR (125 MHz, CDCl_3) δ

(ppm): 169.6, 160.6, 141.7, 136.4, 129.0, 127.4, 126.4, 121.5, 114.7, 86.7, 55.4, 37.4, 37.2, 36.3; HRMS (EI) calcd for C₁₈H₁₈O₃S 314.0977, found 314.0978; HPLC (Chiralpak IC, *i*-propanol/hexane = 20/80, flow rate 1.0 mL/min, λ = 254 nm): t_R (major) = 43.3 min, t_R (minor) = 28.1 min, *ee* = 92%; $[\alpha]^{25}_D$ = -60.3 (*c* = 1.08 in DCM).



(4*S*,6*S*)-6-(2-methoxyphenylthio)-4-phenyltetrahydro-2*H*-pyran-2-one (3ag) (Table 2, entry 15).

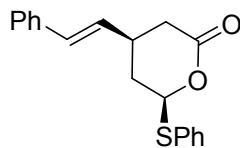
52.7 mg, 84% yield; Colorless oil; ¹H NMR (500 MHz, CDCl₃) δ (ppm): 7.61 (dd, *J* = 7.6, 1.6 Hz, 1H), 7.37 – 7.27 (m, 4H), 7.21 – 7.18 (m, 2H), 6.97 (td, *J* = 7.5, 1.1 Hz, 1H), 6.91 (d, *J* = 8.2 Hz, 1H), 5.87 (dd, *J* = 10.9, 4.3 Hz, 1H), 3.89 (s, 3H), 3.20 (tt, *J* = 12.6, 4.4 Hz, 1H), 2.88 (ddd, *J* = 17.5, 5.0, 2.3 Hz, 1H), 2.63 – 2.56 (m, 2H), 2.05 (ddd, *J* = 14.0, 12.2, 11.0 Hz, 1H); ¹³C NMR (125 MHz, CDCl₃) δ (ppm): 169.6, 158.4, 141.7, 134.4, 130.0, 129.0, 127.4, 126.4, 121.4, 119.9, 111.1, 84.5, 55.9, 37.4, 37.3, 36.0; HRMS (EI) calcd for C₁₈H₁₈O₃S 314.0977, found 314.0979; HPLC (Chiralpak IA, *i*-propanol/hexane = 20/80, flow rate 1.0 mL/min, λ = 254 nm): t_R (major) = 11.1 min, t_R (minor) = 13.1 min, *ee* = 94%; $[\alpha]^{25}_D$ = -67.5 (*c* = 0.96 in DCM).



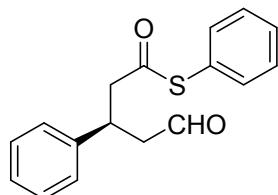
(4*S*,6*S*)-6-(3,4-dimethoxyphenylthio)-4-phenyltetrahydro-2*H*-pyran-2-one (3ah) (Table 2, entry

16). 54.8 mg, 80% yield; Colorless oil; ¹H NMR (500 MHz, CDCl₃) δ (ppm): 7.35 (dd, *J* = 10.2, 4.7 Hz, S11

2H), 7.30 – 7.25 (m, 2H), 7.19 (dd, J = 8.3, 2.1 Hz, 1H), 7.16 – 7.13 (m, 3H), 6.85 (d, J = 8.3 Hz, 1H), 5.63 (dd, J = 11.2, 4.2 Hz, 1H), 3.89 (s, 6H), 3.20 – 3.13 (m, 1H), 2.85 (ddd, J = 17.7, 5.1, 2.3 Hz, 1H), 2.55 – 2.43 (m, 2H), 1.94 (ddd, J = 13.9, 12.3, 11.3 Hz, 1H); ^{13}C NMR (125 MHz, CDCl_3) δ (ppm): 169.6, 150.1, 149.1, 141.6, 129.0, 127.9, 127.4, 126.4, 121.7, 117.7, 111.4, 86.6, 56.1, 55.9, 37.3, 37.21, 36.3; HRMS (EI) calcd for $\text{C}_{19}\text{H}_{20}\text{O}_4\text{S}$ 344.1082, found 344.1081; HPLC (Chiralpak IA, *i*-propanol/hexane = 5/95, flow rate 1.0 mL/min, λ = 254 nm): t_R (major) = 46.4 min, t_R (minor) = 52.4 min, ee = 92%; $[\alpha]^{25}_{\text{D}} = -61.1$ (c = 0.99 in DCM).



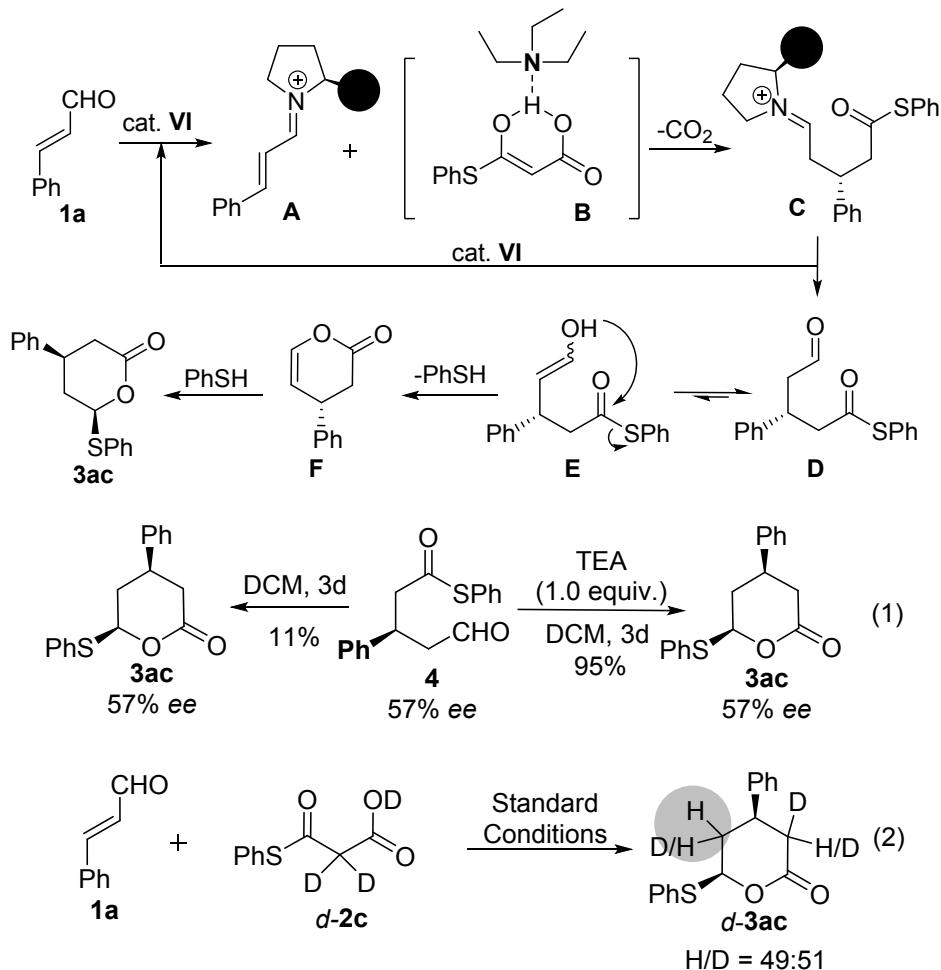
(4*S*,6*S*)-6-(phenylthio)-4-styryltetrahydro-2*H*-pyran-2-one (3lc) (Table 2, entry 17). 46.2 mg, 74% yield; Pale yellow oil; ^1H NMR (500 MHz, CDCl_3) δ (ppm): 7.55 (dd, J = 6.5, 2.9 Hz, 2H), 7.31 – 7.21 (m, 8H), 6.39 (d, J = 15.9 Hz, 1H), 5.97 (dd, J = 15.9, 7.0 Hz, 1H), 5.63 (dd, J = 11.2, 4.0 Hz, 1H), 2.82 – 2.72 (m, 2H), 2.42 (ddd, J = 13.9, 6.1, 3.8 Hz, 1H), 2.29 (dd, J = 17.2, 11.4 Hz, 1H), 1.72 (dt, J = 13.8, 11.5 Hz, 1H); ^{13}C NMR (125 MHz, CDCl_3) δ (ppm): 169.2, 136.3, 133.3, 131.7, 130.8, 129.9, 129.2, 128.7, 128.6, 127.9, 126.2, 86.1, 35.7, 35.1, 35.1; HRMS (EI) calcd for $\text{C}_{19}\text{H}_{18}\text{O}_2\text{S}$ 310.1028, found 310.1035; HPLC (Chiralpak IC, *i*-propanol/hexane = 20/80, flow rate 1.0 mL/min, λ = 254 nm): t_R (major) = 36.6 min, t_R (minor) = 29.6 min, ee = 80%; $[\alpha]^{25}_{\text{D}} = -48.2$ (c = 0.78 in DCM).



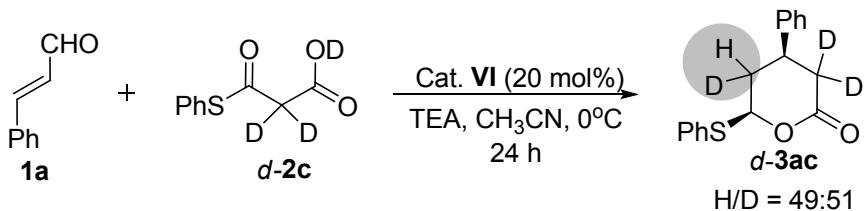
(R)-S-phenyl 5-oxo-3-phenylpentanethioate (4). Colorless oil; ^1H NMR (500 MHz, CDCl_3) δ (ppm): 9.74 (s, 1H), 7.47 – 7.45 (m, 3H), 7.42 – 7.38 (m, 4H), 7.34 – 7.31 (m, 3H), 3.95 – 3.86 (m, 1H), 3.09 (d, J = 7.2 Hz, 2H), 3.00 – 2.89 (m, 2H); ^{13}C NMR (125 MHz, CDCl_3) δ (ppm): 200.4, 195.6, 141.8, 134.4, 129.5, 129.2, 128.8, 127.4, 127.2, 49.5, 49.0, 36.7; HRMS (EI) calcd for $\text{C}_{17}\text{H}_{16}\text{O}_2\text{S}$ 284.0871, found 284.0878.

3. Proposed Mechanism

Our proposed mechanism is illustrated in Scheme below. Initially, malonic half-thioester **2a** reacts with triethyl amine to form the intermediate **B** and follow to attack the iminium intermediate **A**. A subsequent decarboxylation triggers the formation of the Michael addition intermediate **C**. A hydrolysis of intermediate **C** would lead to a recycle of cat.**VI** and generates the intermediate **D**. The intermediate **D** undergoes a sequence of tautomerization, cyclization and nucleophilic addition to afford the product **3ac**. As proof of this mechanism, we set out to investigate the key intermediate **4**. As indicated in eqn. (2), the synthesized compound **4** was found to be an active intermediate and could generate **3ac** under weak base or base-free condition (95% and 11% yield, respectively) and gave no loss of *ee*. Meanwhile, we found that this process did not require Cat. **IV**. This result suggests that the compound **4** may involve in this transformation as a potential key intermediate. To further understand this mechanism, we then investigated the proton transfer in the process of **3ac** formation. we conducted an isotopic experiment using deuterated half-thioester *d*-**2c** to react with cinnamaldehyde **1a** under standard conditions. ^1H NMR analysis of *d*-**3ac** demonstrated the isotope labelling of two protons located at α -position of carbonyl group is ab. 49:51 (eqn. (2)). It suggests that the water (generated from the condensation of cinnamaldehyde **1a** and cat. **VI**), functionalized as a potential proton source, is not involved in Michael addition step.



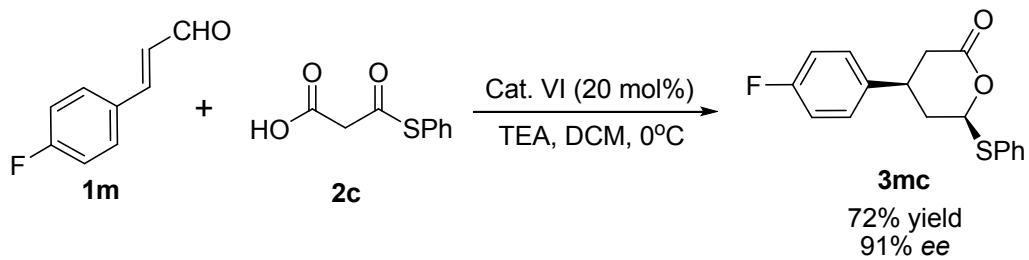
490.6 mg (2.5 mmol) of malonic acid half thioester (MAHT) **2c** was placed into a 50 mL flask and 5.0 mL of acetonitrile was added. When the MAHT **2c** has dissolved, 6.0 mL D₂O was added and the solution was stirred for 6 hours under nitrogen. The solvent was removed under vacuum and the process was repeated two more times to yield 473.2 mg (95% yield) of the product *d*-**2c**. ^[1]¹H NMR (500 MHz, CDCl₃) δ (ppm): 7.46 – 7.42 (m, 5H), 3.70 (s, 0.2H); ¹³C NMR (125 MHz, CDCl₃) δ (ppm): 189.9, 170.5, 134.4, 130.0, 129.4, 126.4, 48.0 (t, 1C, *J* = 20.5 Hz).



To a solution of cinnamaldehyde **1a** (66.1 mg, 62.9 uL, 0.5 mmol,) in CH₃CN (2.5 mL) was added catalyst **VI** (64.0 mg, 0.1 mmol) at 0 °C. After 40 min, *d*-**2c** (249.0 mg, 1.25 mmol) was added in one portion. Then, TEA (126.2 mg, 173.9 uL, 1.25 mmol) was added dropwise at 0 °C. The resulting reaction mixture was kept stirring at 0°C for 24 h. The crude product was purified by silica gel flash chromatography, eluted by hexane/EtOAc = 10:1 to afford the desired product *d*-**3ac** as white solid (89.1 mg, 62% yield). ¹H NMR (500 MHz, CDCl₃) δ(ppm): 7.60 (dd, *J* = 6.4, 3.2 Hz, 2H), 7.40 – 7.32 (m, 5H), 7.28 (t, *J* = 7.4 Hz, 1H), 7.16 (d, *J* = 7.6 Hz, 2H), 5.74 (dd, *J* = 11.2, 4.0 Hz, 1H), 3.23 – 3.14 (m, 1H), 2.88 (dd, *J* = 17.7, 5.0 Hz, 0.49H), 2.59 – 2.49 (m, 1H), 2.04 – 1.97 (m, 0.49H); ¹³C NMR (125 MHz, CDCl₃) δ (ppm): 169.5, 141.5, 133.3, 131.8, 129.2, 129.1, 128.6, 127.5, 126.4, 86.3, 86.3, 86.2, 37.4, 37.3, 37.2, 36.3.

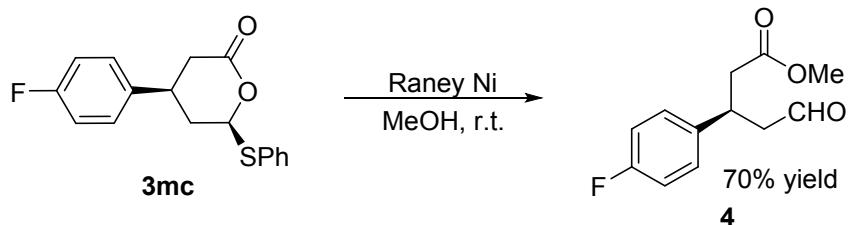
4. Application

Synthesis of (-)-Paroxetine

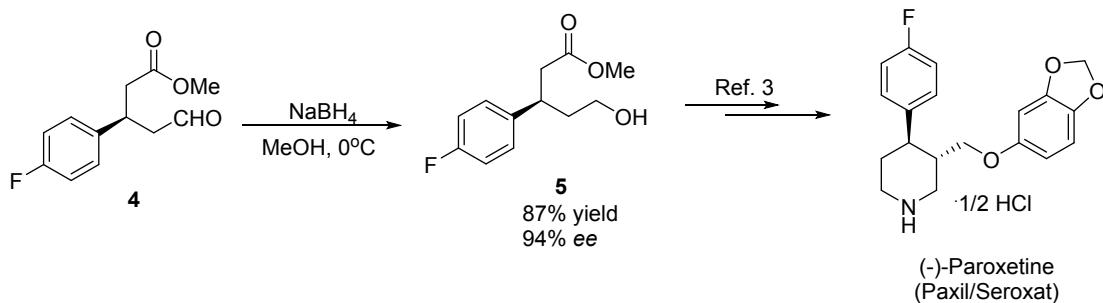


To a solution of 4-fluorocinnamaldehyde **1m** (300.1 mg, 2.0 mmol,) in DCM (5.0 mL) was added catalyst **VI** (238.8 mg, 0.4 mmol) at 0 °C. After 40 min, malonic acid half thioester **2c** (981.1 mg, 5.0 mmol) was added in one portion. Then, TEA (695.6 uL, 5.0 mmol) in 5.0 mL DCM was added

dropwise *via* syringe at 0 °C. The resulting reaction mixture was kept stirring at 0°C for 24 h. The crude product was purified by silica gel flash chromatography, eluted by hexane/EtOAc = 10:1 to afford the desired product **3mc** as a white solid (436.2 mg, 72% yield, 91% *ee*). ¹H NMR (500 MHz, CDCl₃) δ = 7.59 (dd, *J* = 6.2, 2.8 Hz, 2H), 7.39 – 7.33 (m, 3H), 7.13 (dd, *J* = 8.5, 5.3 Hz, 2H), 7.04 (t, *J* = 8.6 Hz, 2H), 5.72 (dd, *J* = 11.2, 4.1 Hz, 1H), 3.18 (ddd, *J* = 12.3, 8.4, 4.5 Hz, 1H), 2.87 (ddd, *J* = 17.6, 5.0, 2.1 Hz, 1H), 2.57 – 2.46 (m, 2H), 1.97 (dd, *J* = 25.7, 11.8 Hz, 1H); ¹³C NMR (125 MHz, CDCl₃) δ = 169.2, 163.0, 161.0, 137.3, 137.3, 133.4, 131.7, 129.2, 128.7, 128.0, 127.9, 116.1, 115.9, 86.2, 37.4, 36.8, 36.4; HRMS (EI) calcd for C₁₇H₁₅FO₂S 302.0777, found 302.0771; HPLC (Chiralpak IC, *i*-propanol/hexane = 20/80, flow rate 1.0 mL/min, λ = 254 nm): *t*_R (minor) = 23.4 min, *t*_R (major) = 25.8 min, *ee* = 91%; [α]_D²⁵ = -78.2 (*c* = 1.51 in DCM).



Compound **3mc** (302.36mg, 1.0 mmol) was added to a suspension of Raney Ni (0.3 g) in absolute MeOH (5.0 mL). The mixture was stirred for 10 hours at room temperature, before the Raney Ni was removed by filtration through a celite pad. Then the catalyst was washed with absolute MeOH (3 × 10 mL) and the filtrate was concentrated under reduced pressure. The crude product was purified by silica gel column chromatography (hexane–EtOAc = 6:1) to afford the compound **4** as a colorless oil (156.1 mg, 70% yield). ¹H NMR (300 MHz, CDCl₃) δ = 9.64 (t, *J* = 1.5 Hz, 1H), 7.23 – 7.12 (m, 2H), 7.03 – 6.94 (m, 2H), 3.78 – 3.68 (m, 1H), 3.58 (s, 3H), 2.88 – 2.71 (m, 2H), 2.71 – 2.56 (m, 2H); ¹³C NMR (75 MHz, CDCl₃) δ = 200.3, 171.8, 163.3, 160.0, 138.1, 138.1, 128.8, 128.7, 115.7, 115.4, 51.6, 49.4, 40.6, 35.3; HRMS (EI) calcd for C₁₂H₁₃FO₃ 224.0849, found 224.0841.



Sodium borohydride (11.3 mg, 0.3 mmol) was added into a solution of compound **4** (44.8 mg, 0.20 mmol) in MeOH (2.0 mL) at 0°C. The reaction mixture was stirred at 0°C for 10 min. Then water (1 mL) was added to quench this reaction. The mixture was extracted with CH₂Cl₂ and the combined organic layers were dried with MgSO₄. Filtration and evaporation of the solvent gave a residue which was purified by silica gel column chromatography (hexane–EtOAc = 3:1) to afford the product **5** as a colorless oil (39.4 mg, 87% yield, 94% *ee*). ¹H NMR (500 MHz, CD₃OD) δ = 7.25 – 7.22 (m, 2H), 7.02 – 6.99 (m, 2H), 3.54 (s, 3H), 3.42 (ddd, *J* = 11.0, 6.9, 5.3 Hz, 1H), 3.37 – 3.32 (m, 1H), 3.31 – 3.24 (m, 1H), 2.70 (dd, *J* = 15.2, 6.4 Hz, 1H), 2.59 (dd, *J* = 15.2, 9.0 Hz, 1H), 1.94 – 1.87 (m, 1H), 1.82 – 1.76 (m, 1H); ¹³C NMR (125 MHz, CD₃OD) δ = 174.3, 164.0, 162.1, 140.8, 140.8, 130.3, 130.3, 116.1, 116.0, 60.5, 51.9, 42.3, 39.9, 39.3; HRMS (EI) calcd for C₁₂H₁₅FO₃ 226.1005, found 226.0999; HPLC (Chiralpak OB-H, *i*-propanol/hexane = 10/90, flow rate 0.8 mL/min, λ = 254 nm): *t*_R (major) = 11.1 min, *t*_R (minor) = 14.1 min, *ee* = 94%; [α]²⁵_D = -16.0 (*c* = 1.06 in DCM).

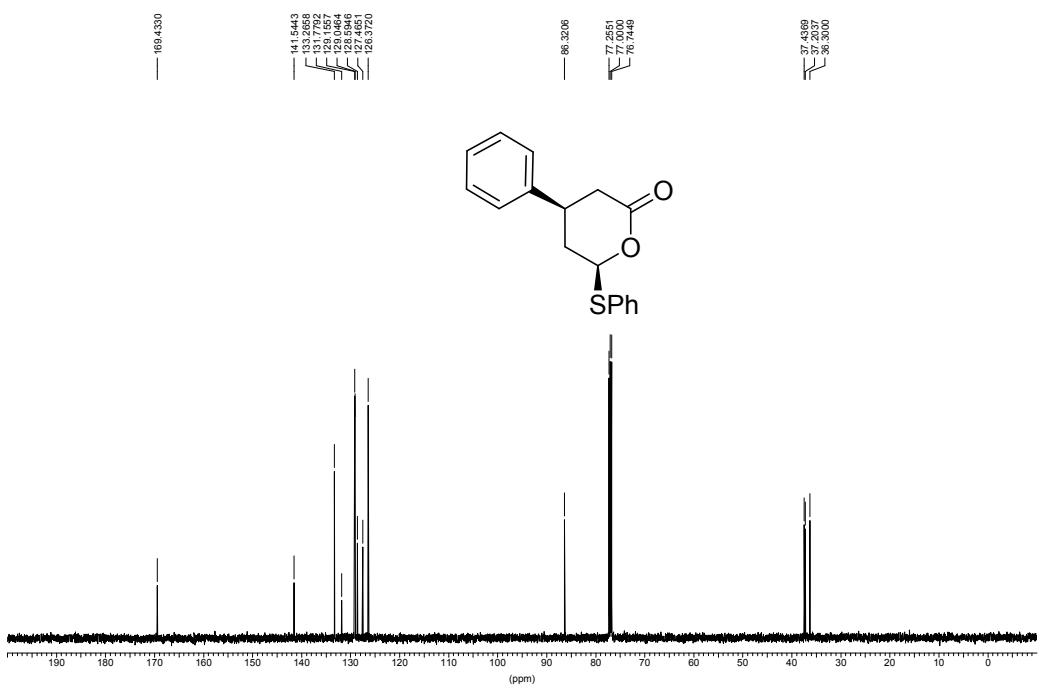
Reference:

- [1] K. C. Fortner, M. D. Shair *J. Am. Chem. Soc.* **2007**, *129*, 1032.
- [2] a) H. Shimizu, S. Fukuda, S. Sugiyama, T. Satoh *Synthesis* **2009**, *8*, 1323–1335; b) M. Ochiai, K. Nishide, M. Node, E. Fujita *Chem. Lett.* **1981**, 283–284.
- [3] M. S. Yu, I. Lantos, Z.-Q. Peng, J. Yu, T. Cacchio *Tetrahedron Lett.* **2000**, *41*, 5647–5651.

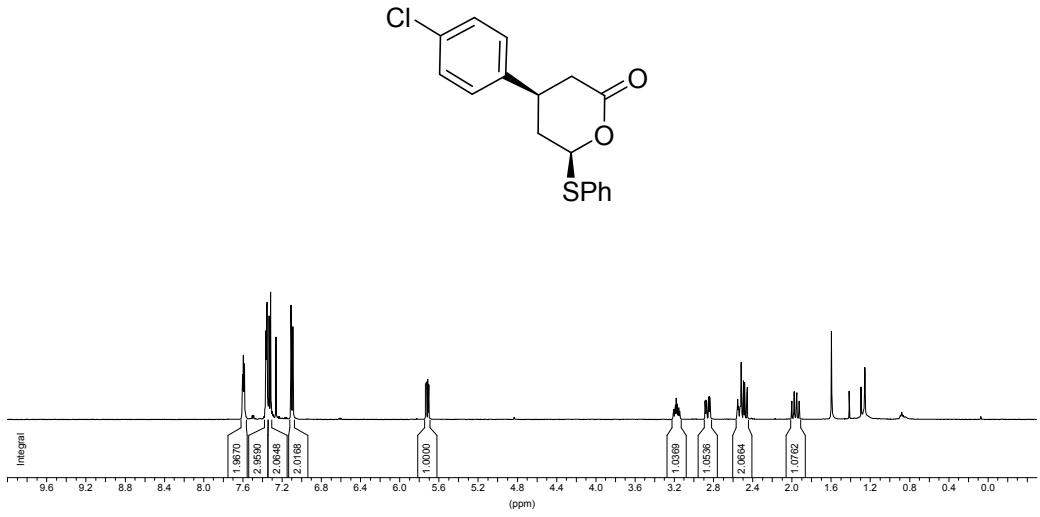
5. HPLC and NMR Spectra

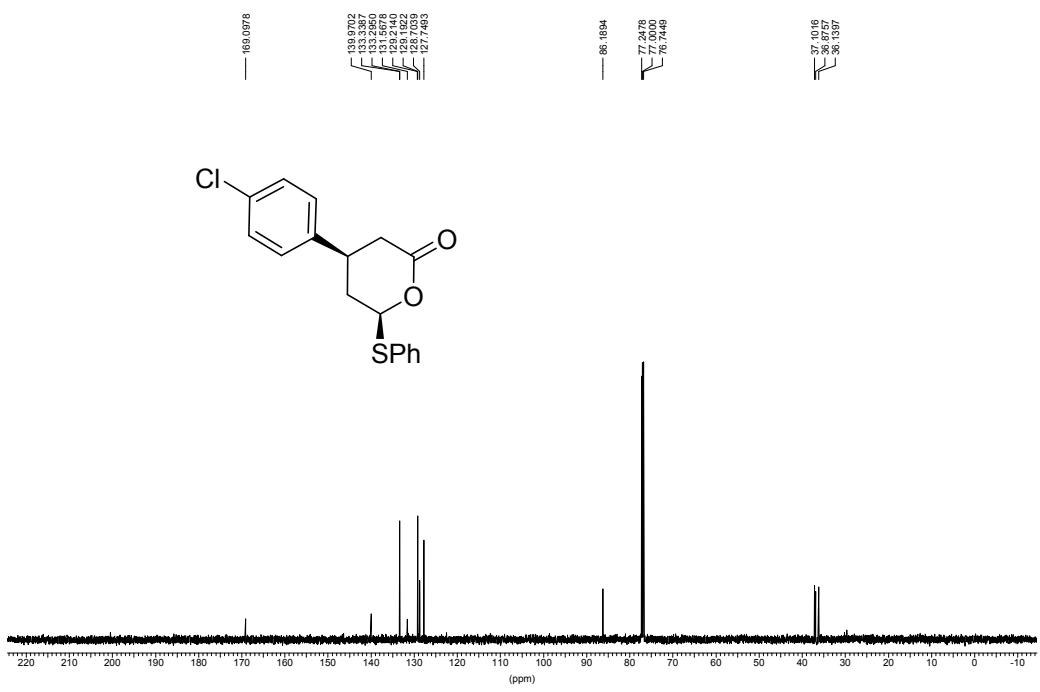
Compound 3ac





Compound 3bc

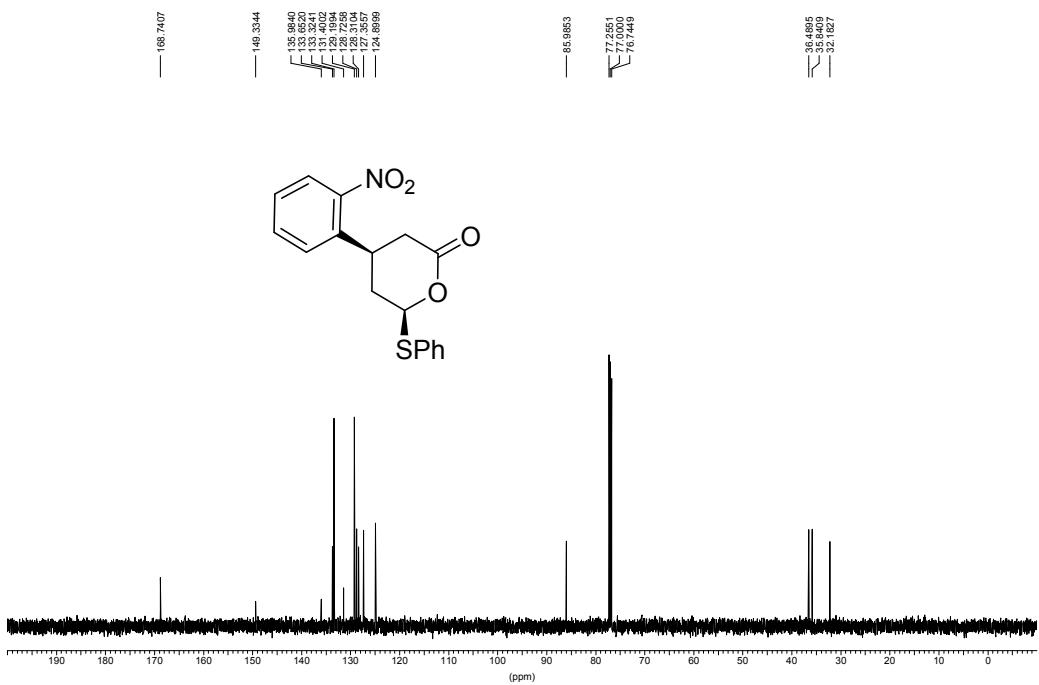
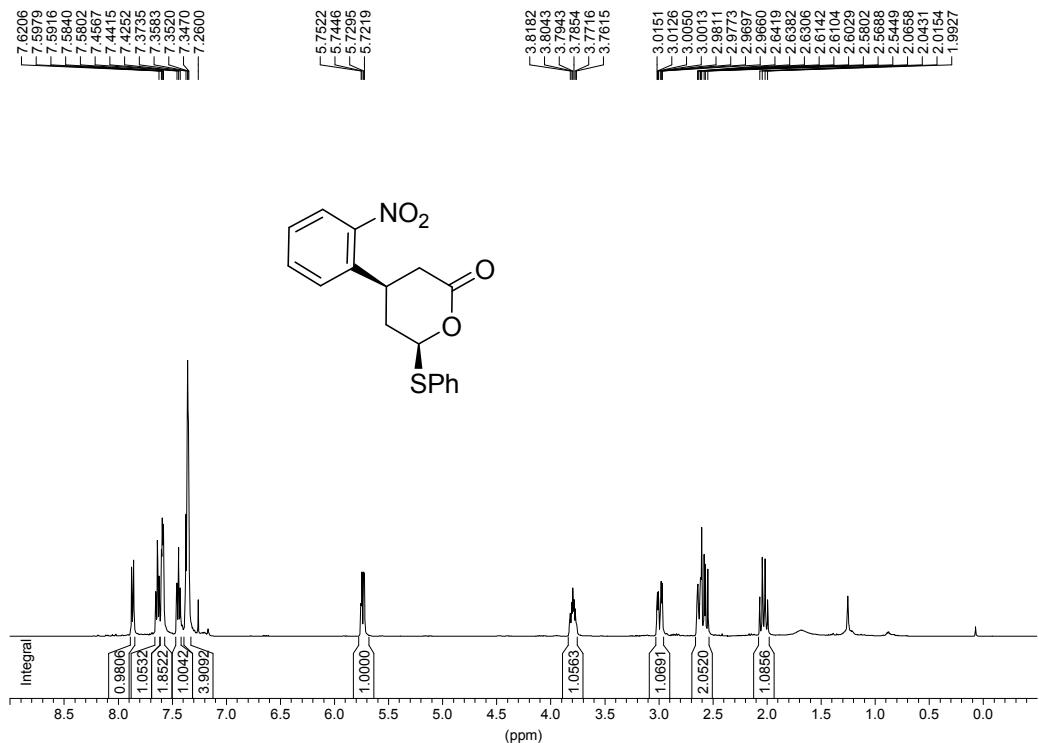




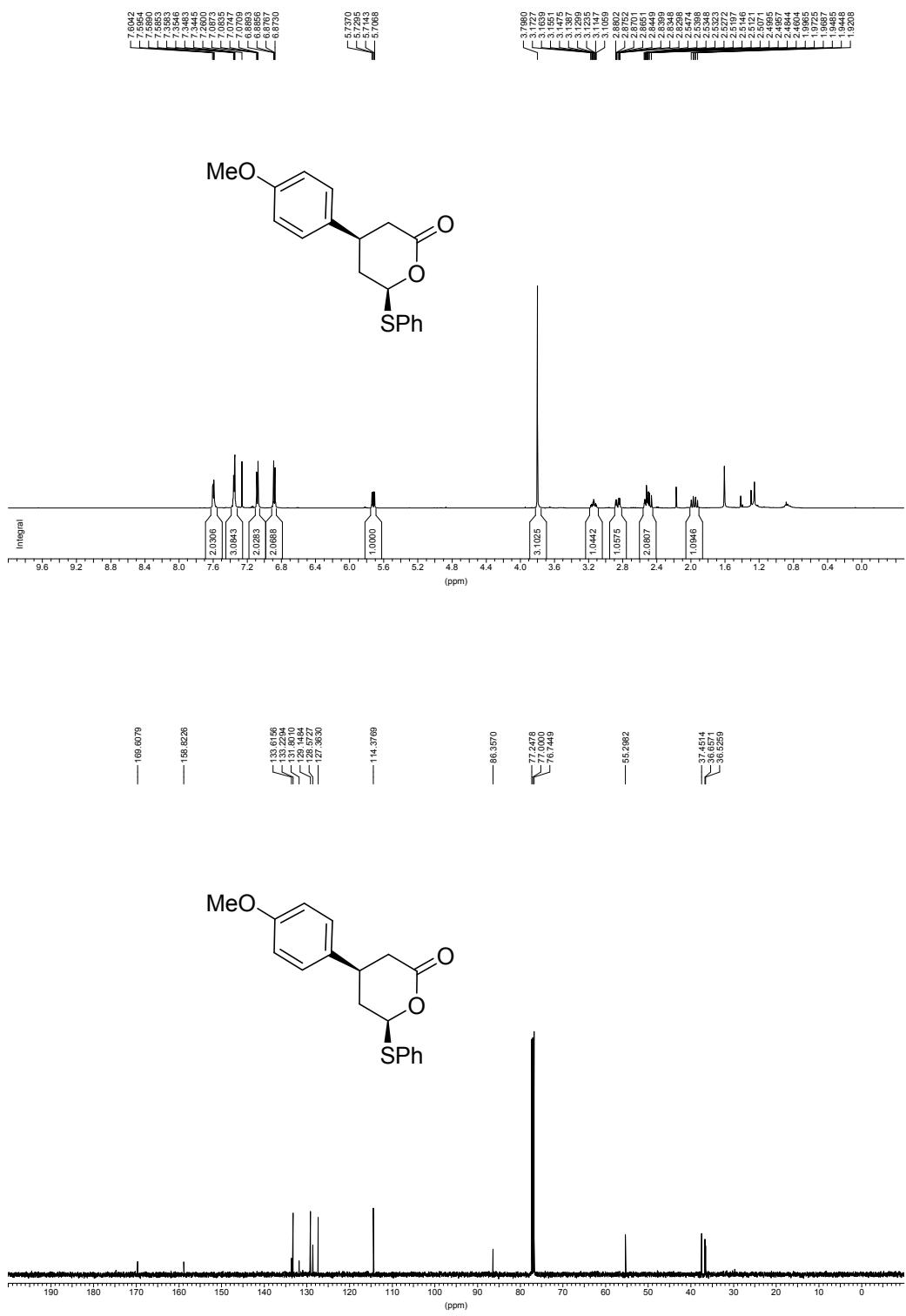
Compound 3cc



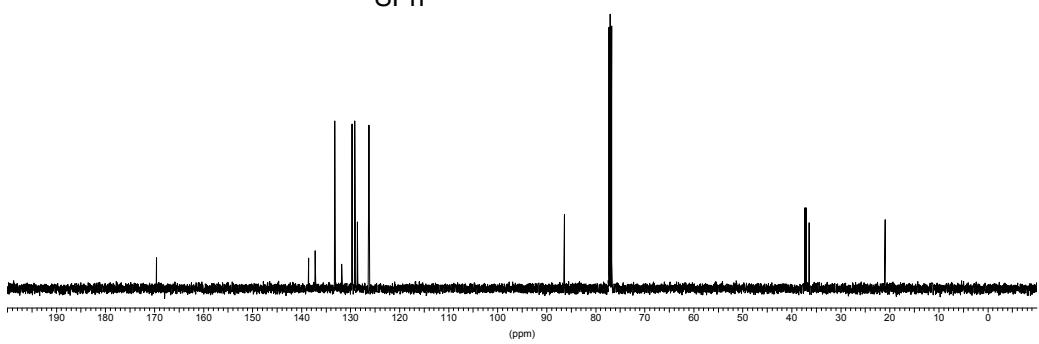
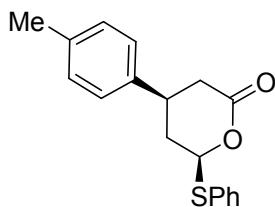
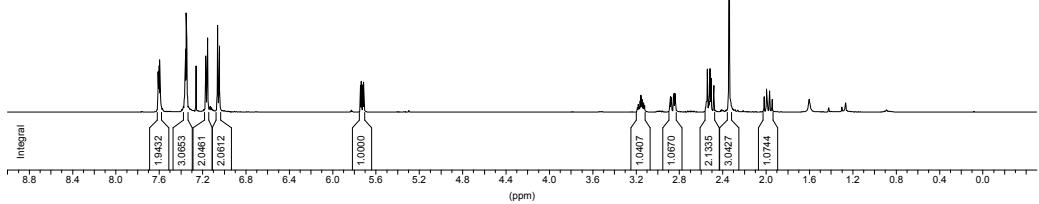
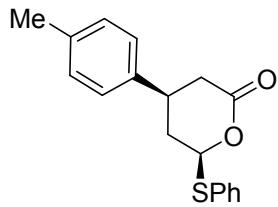
Compound 3dc



Compound 3ec



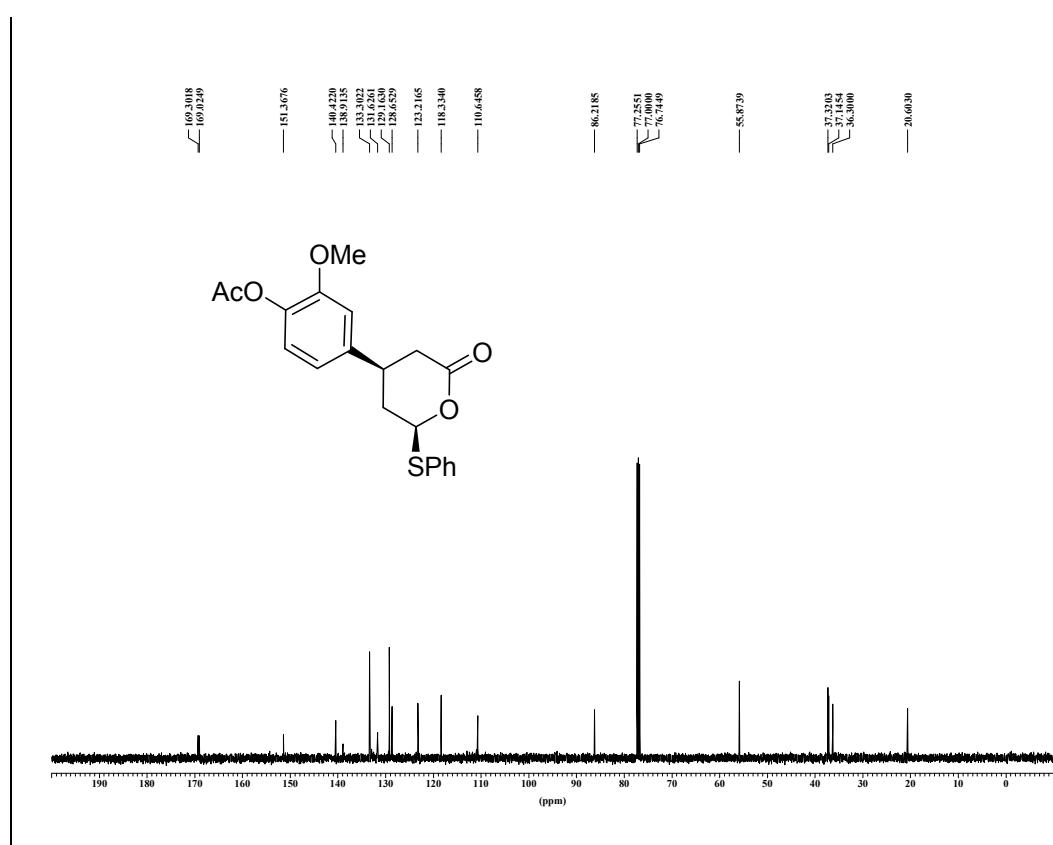
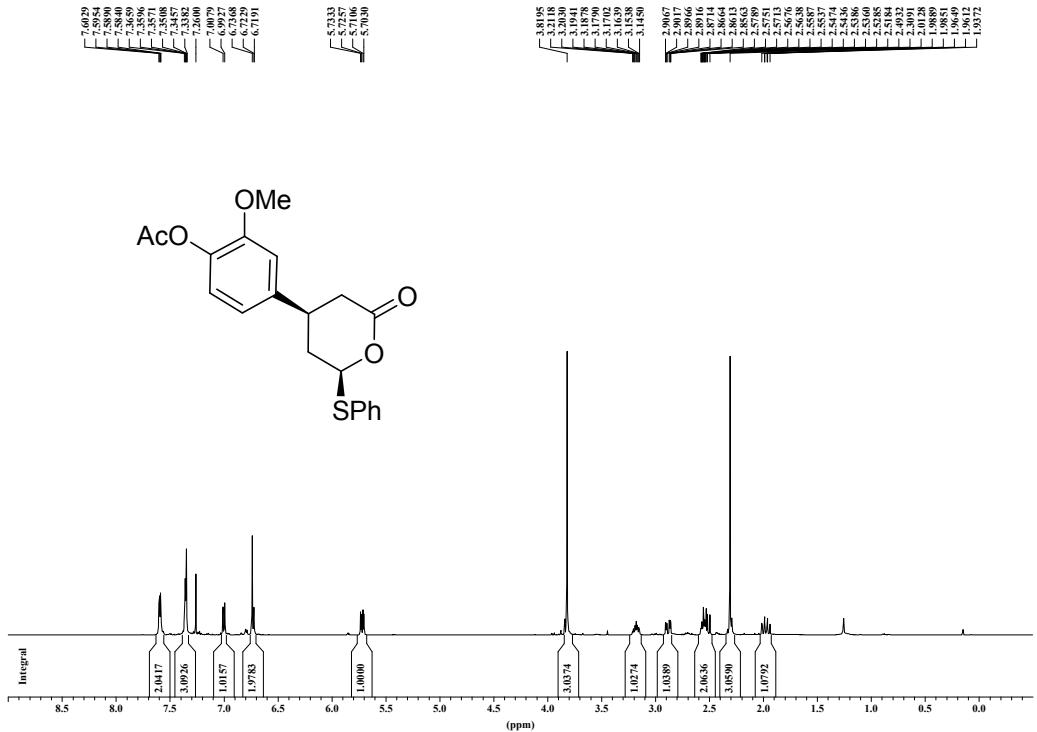
Compound 3fc



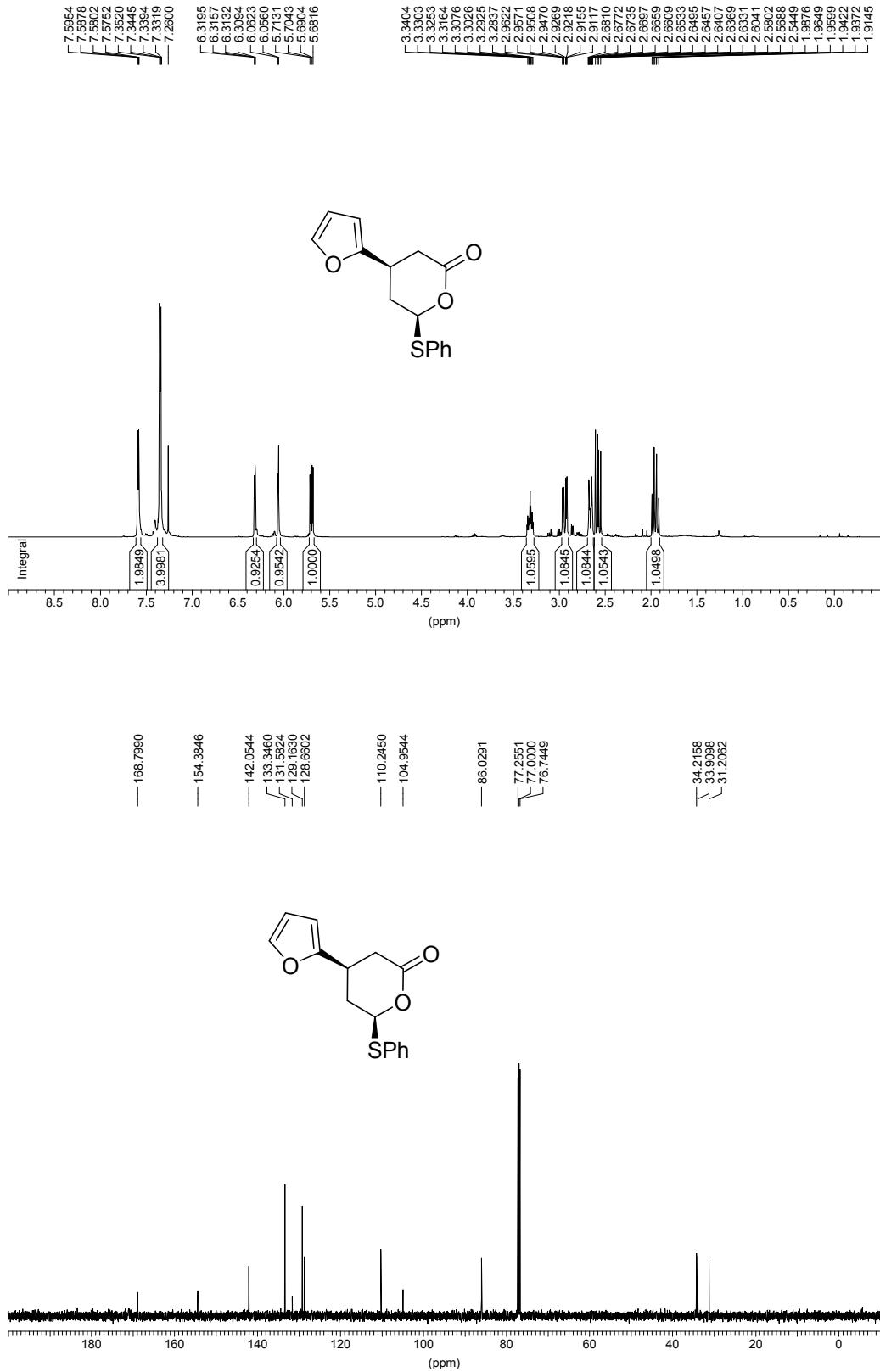
Compound 3gc



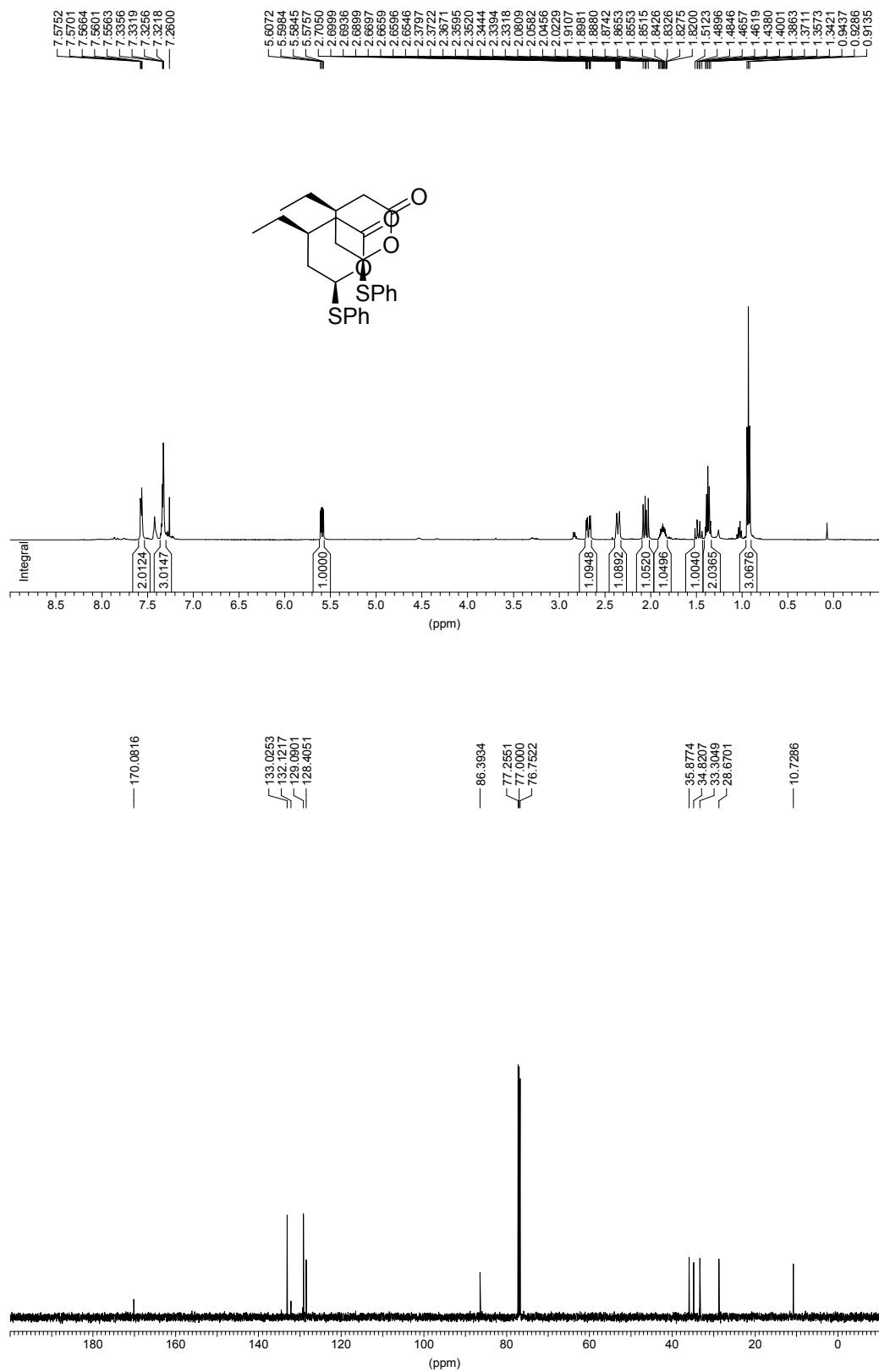
Compound 3hc



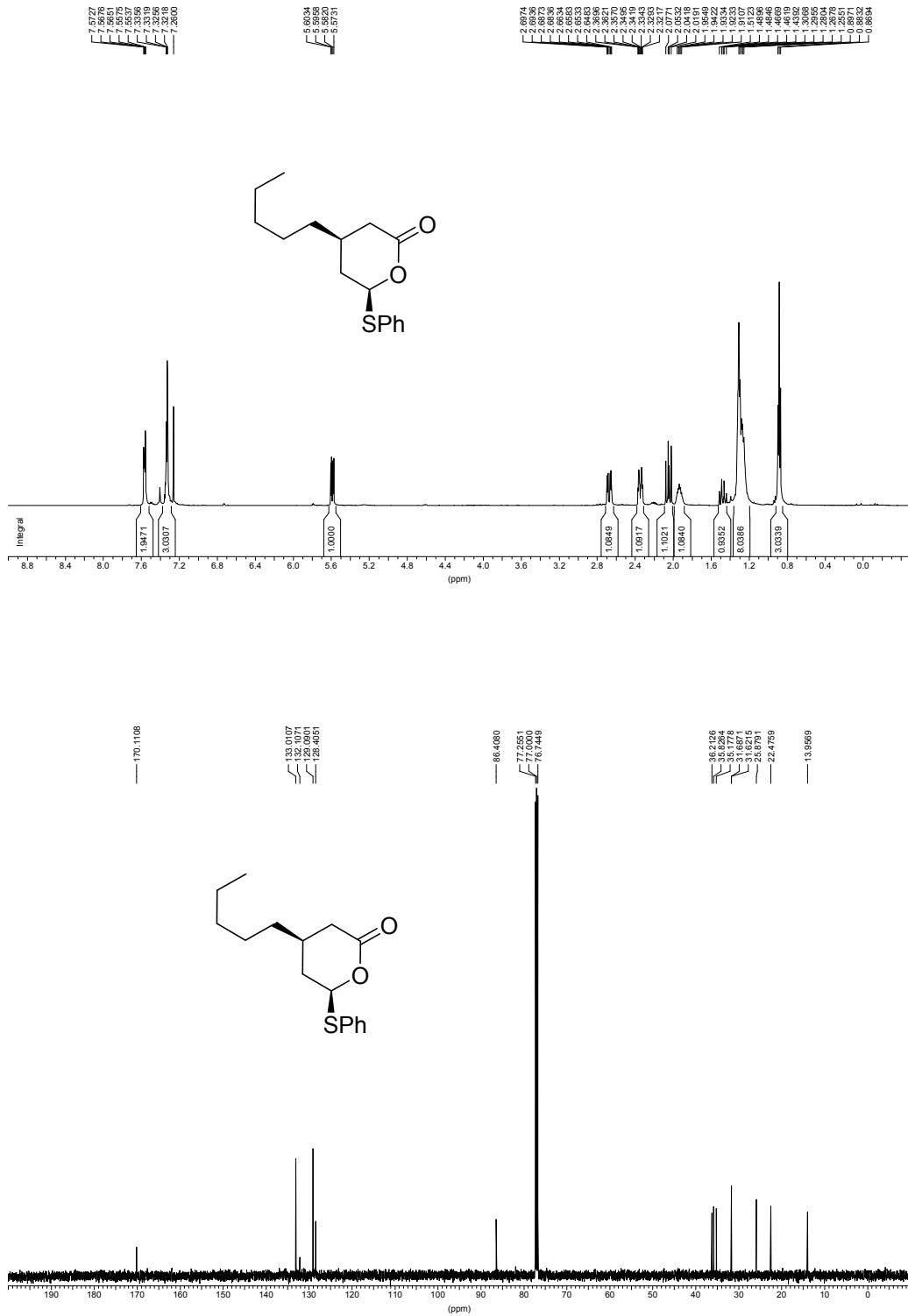
Compound 3ic



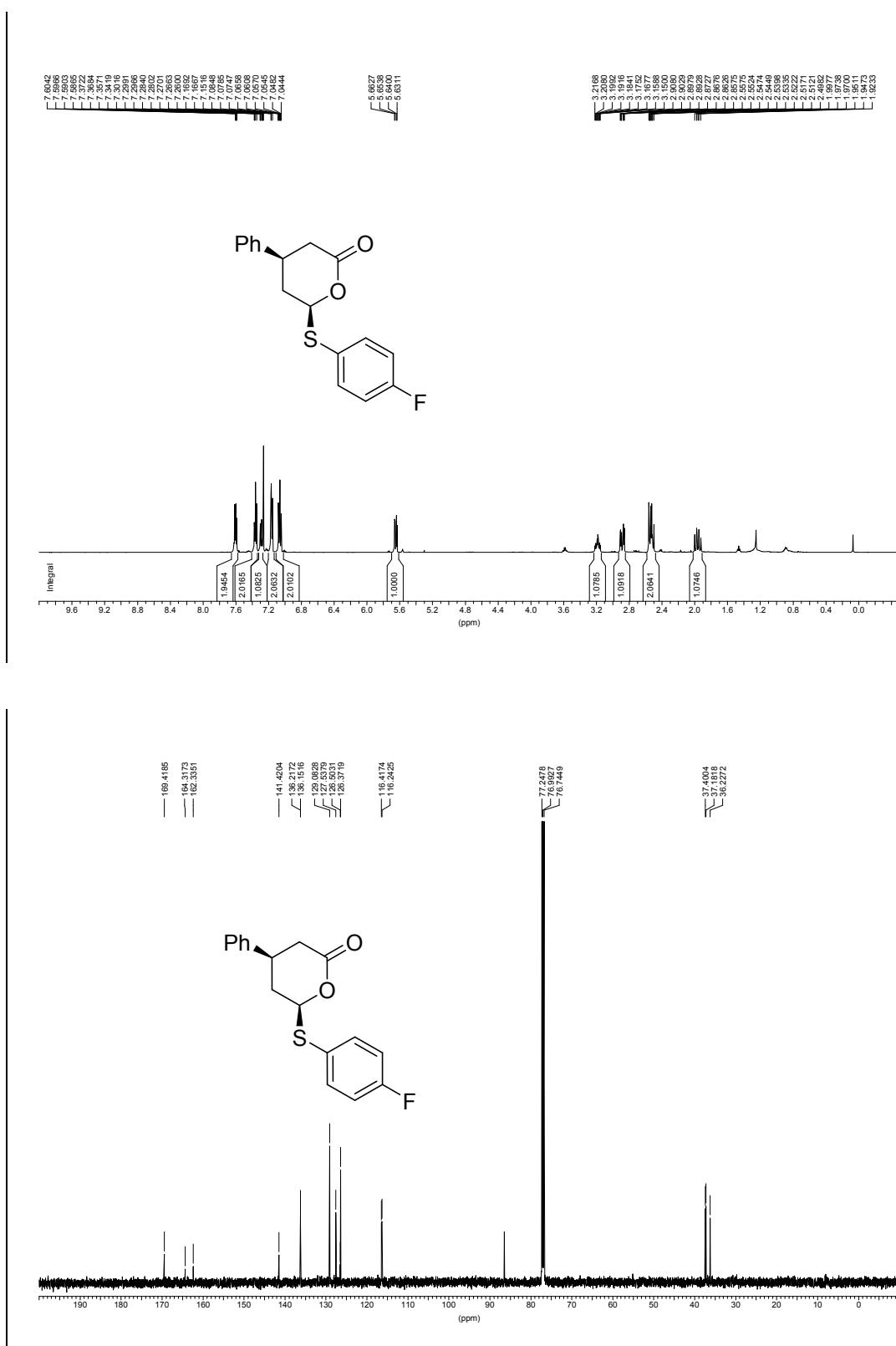
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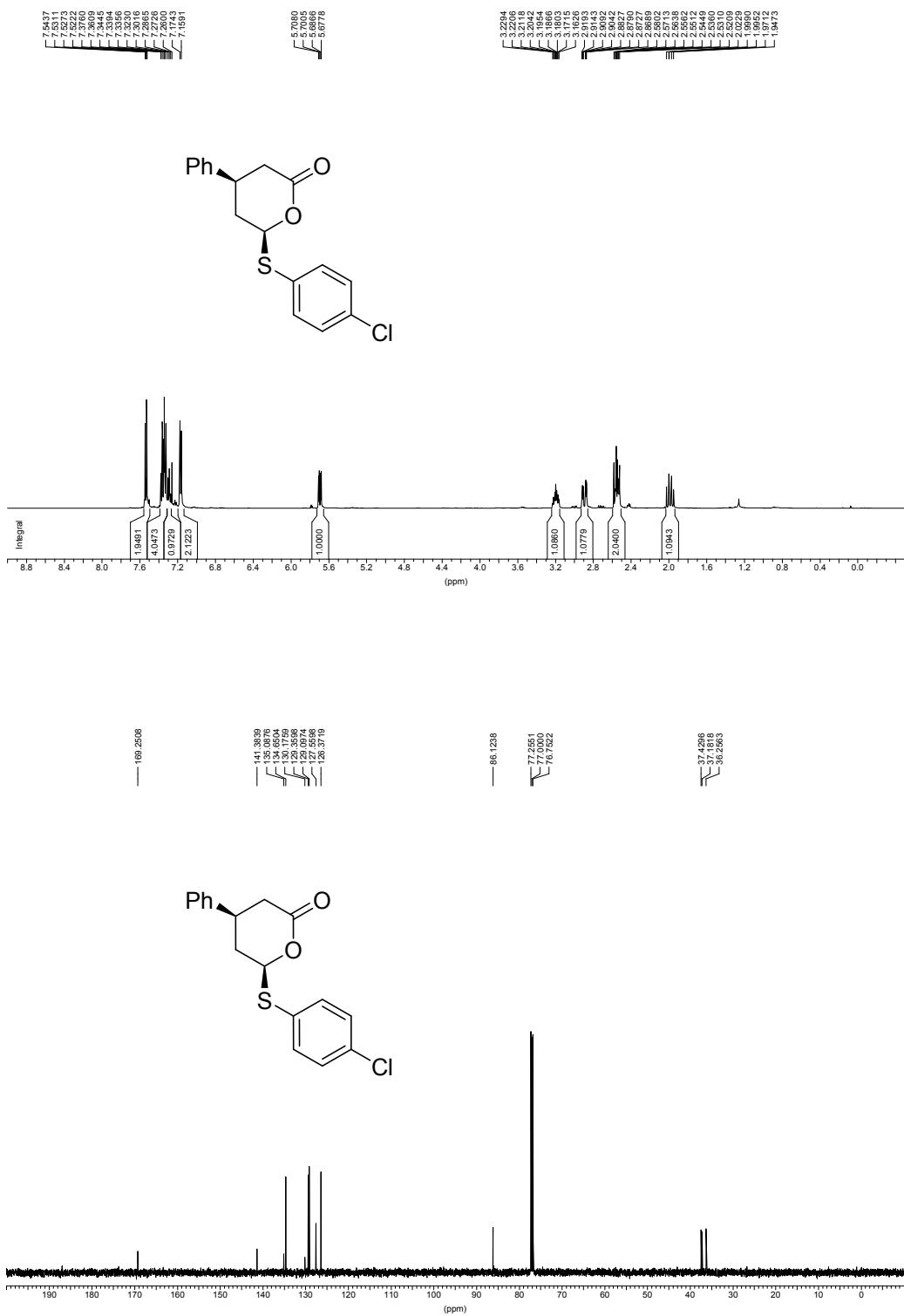
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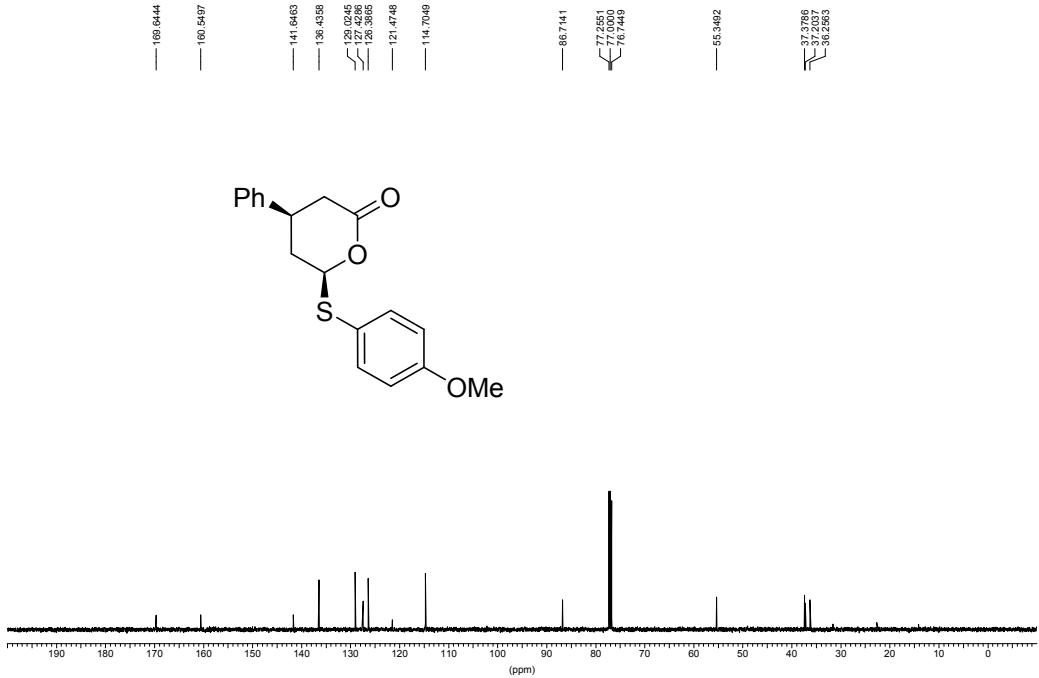
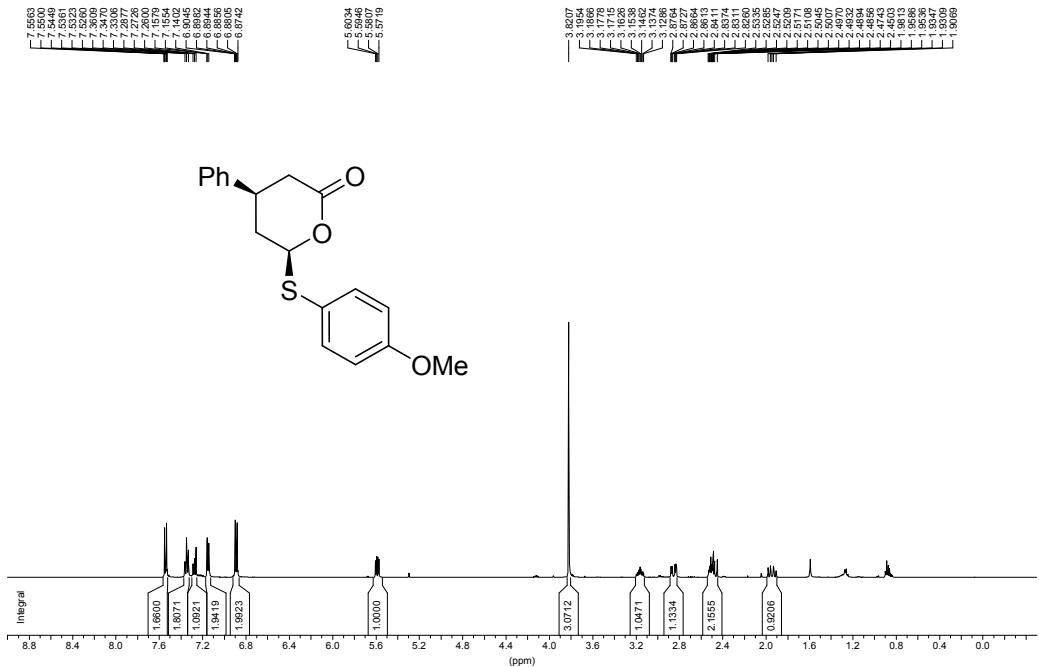
Compound 3ad



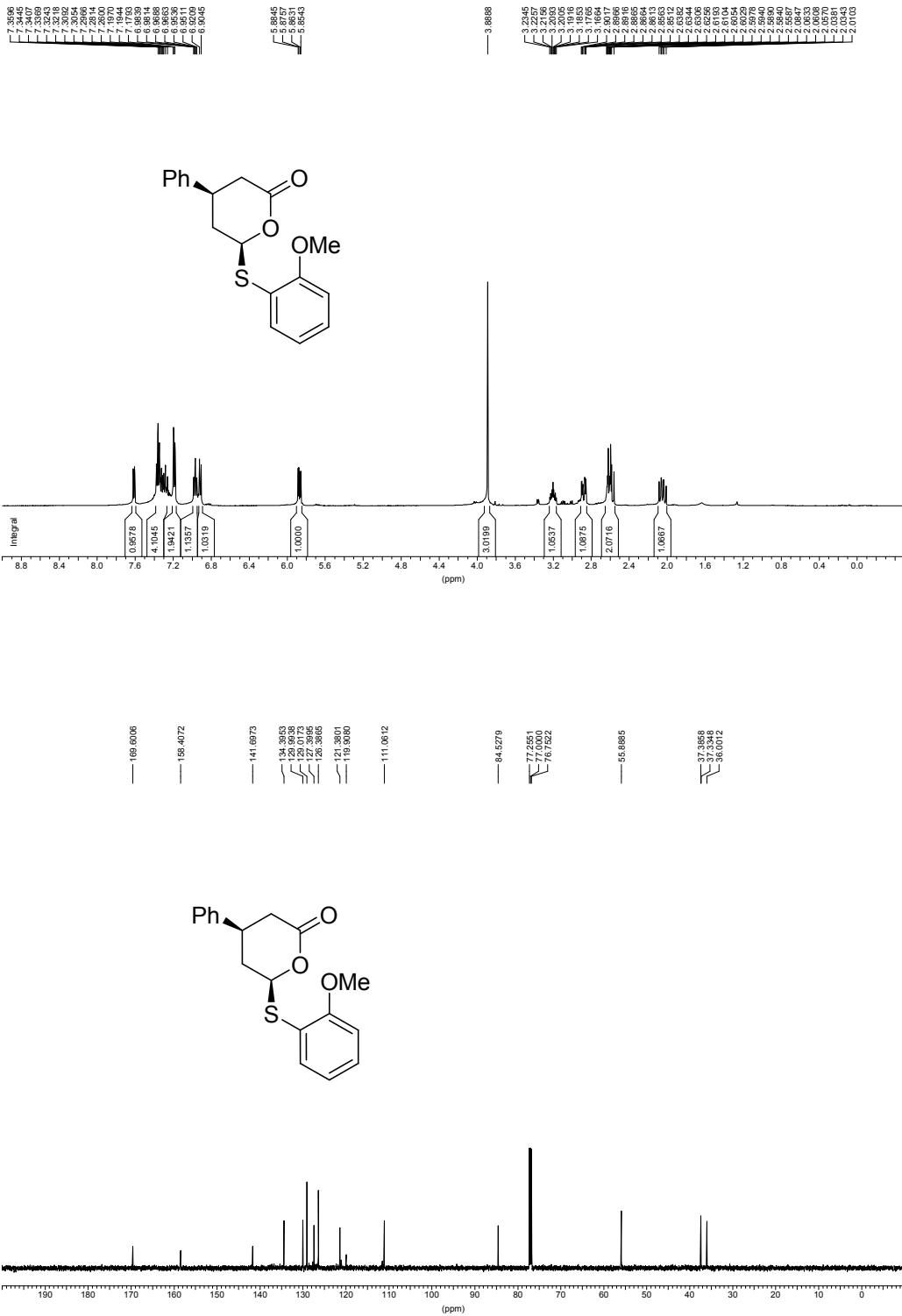
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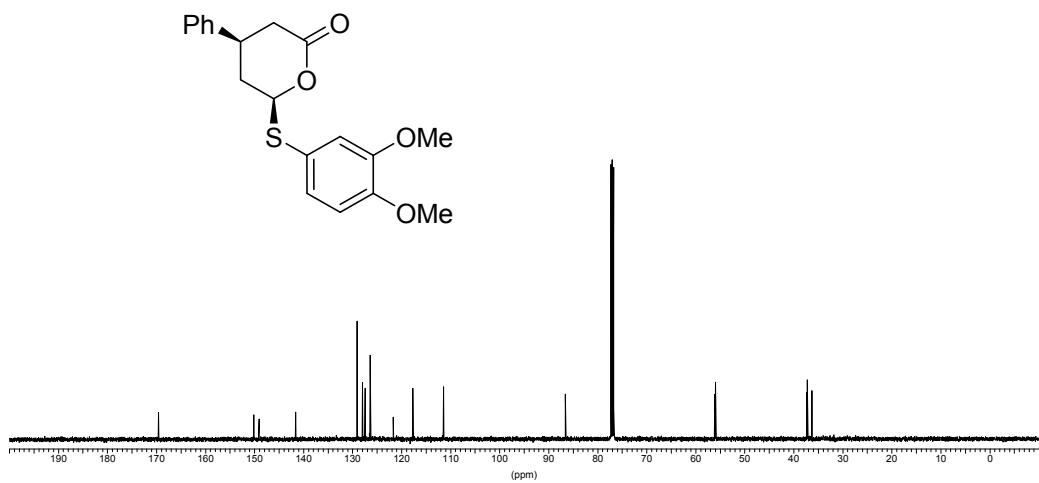
Compound 3af



Compound 3ag



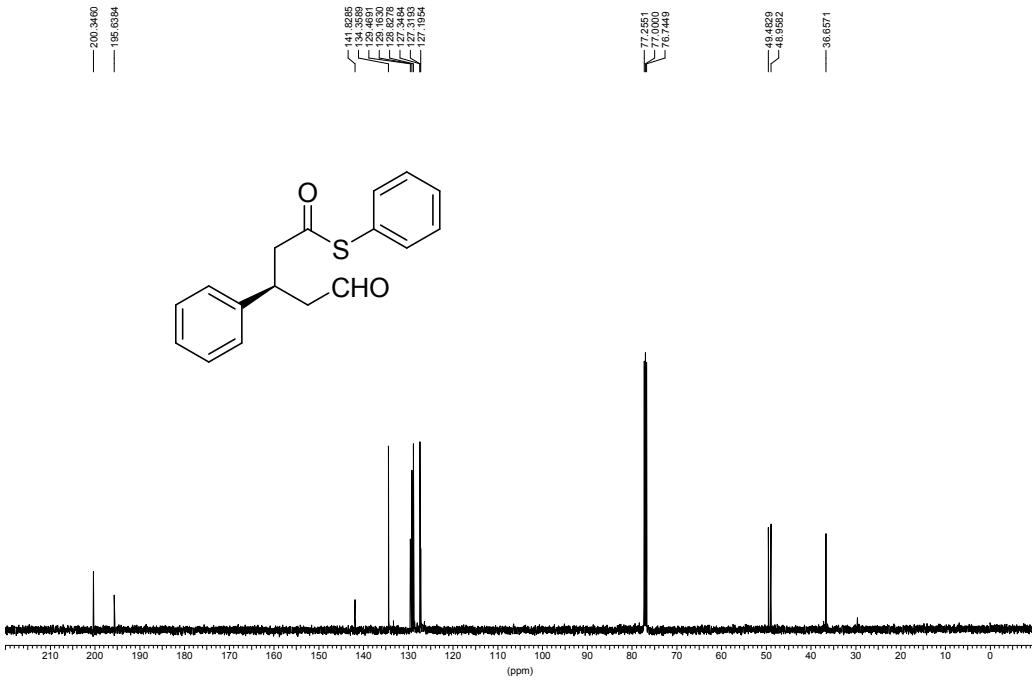
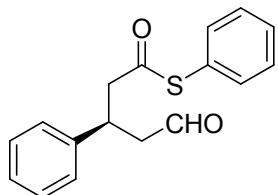
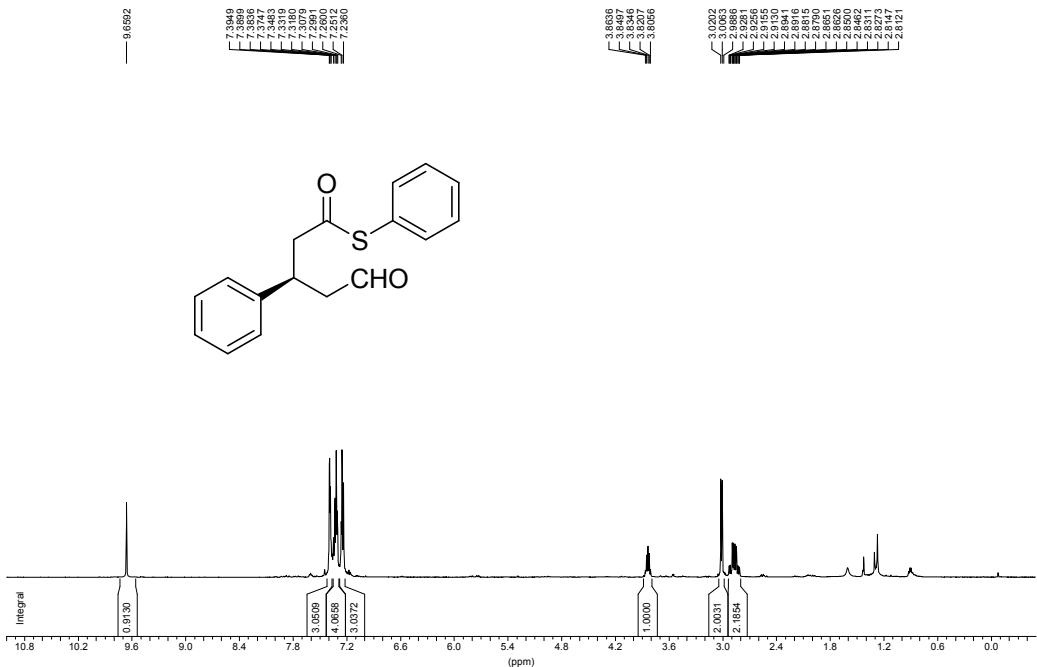
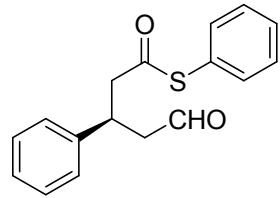
Compound 3ah



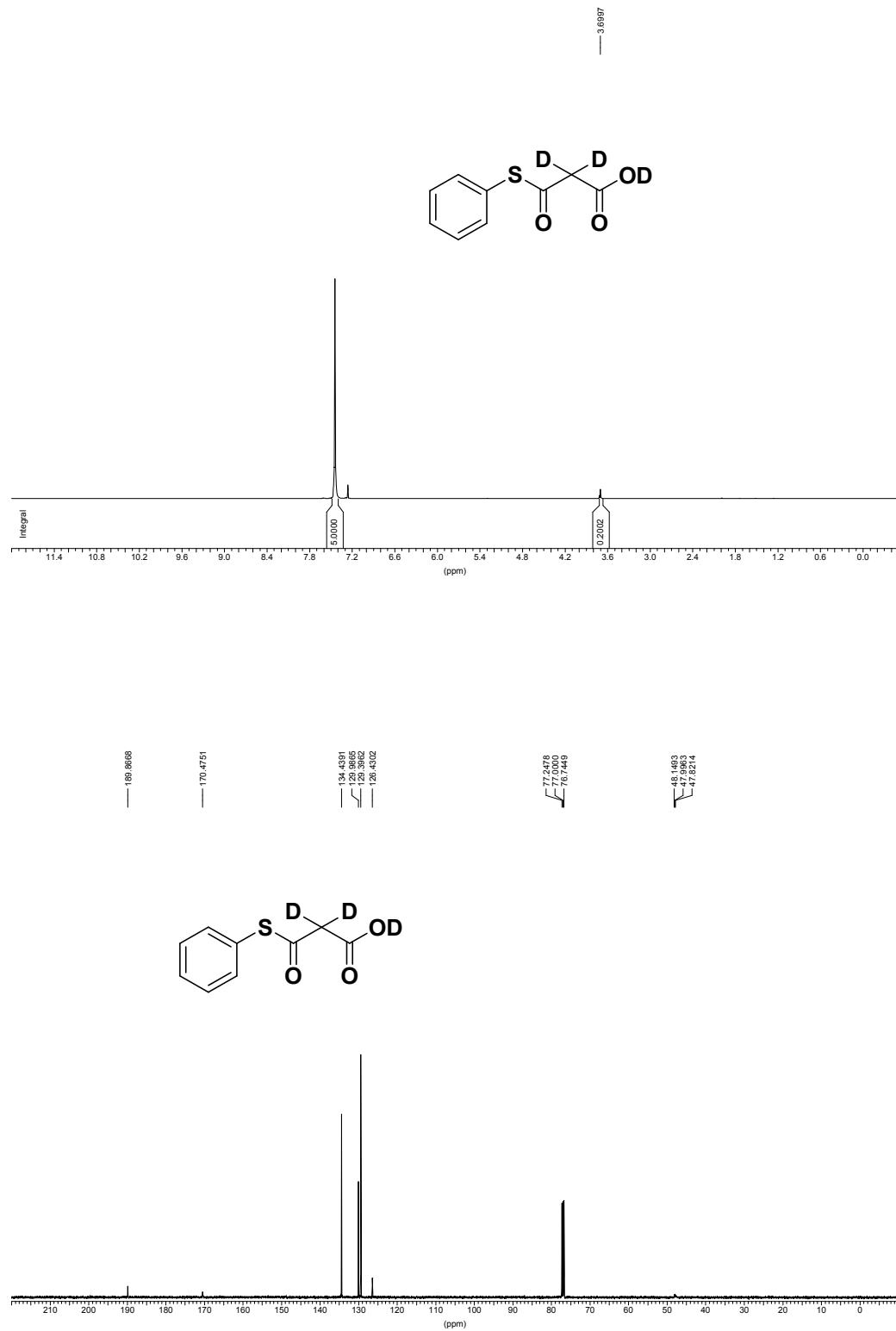
Compound 3lc



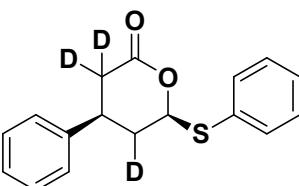
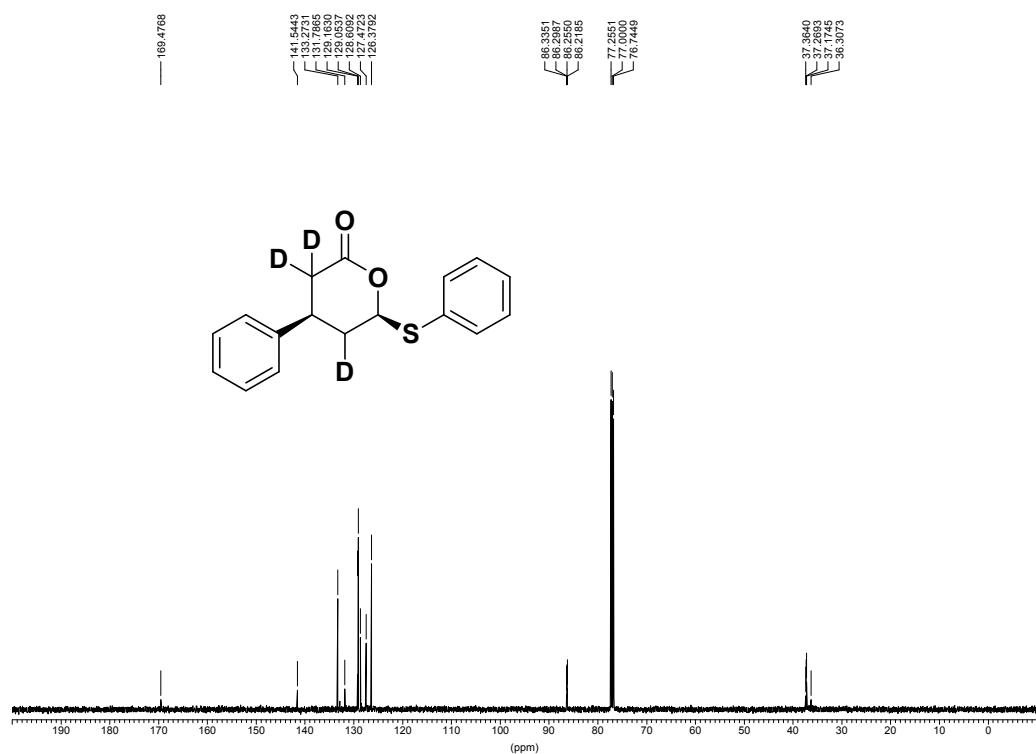
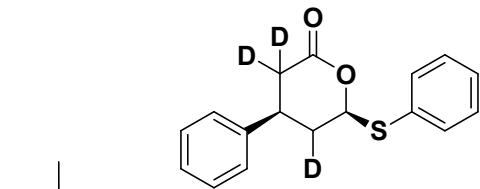
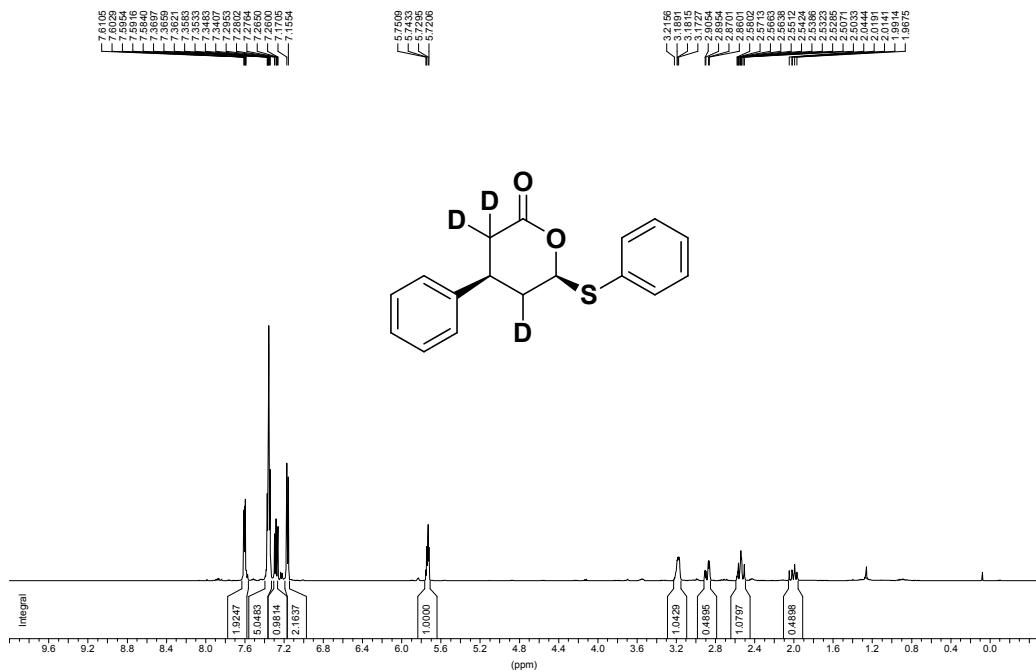
Compound 4



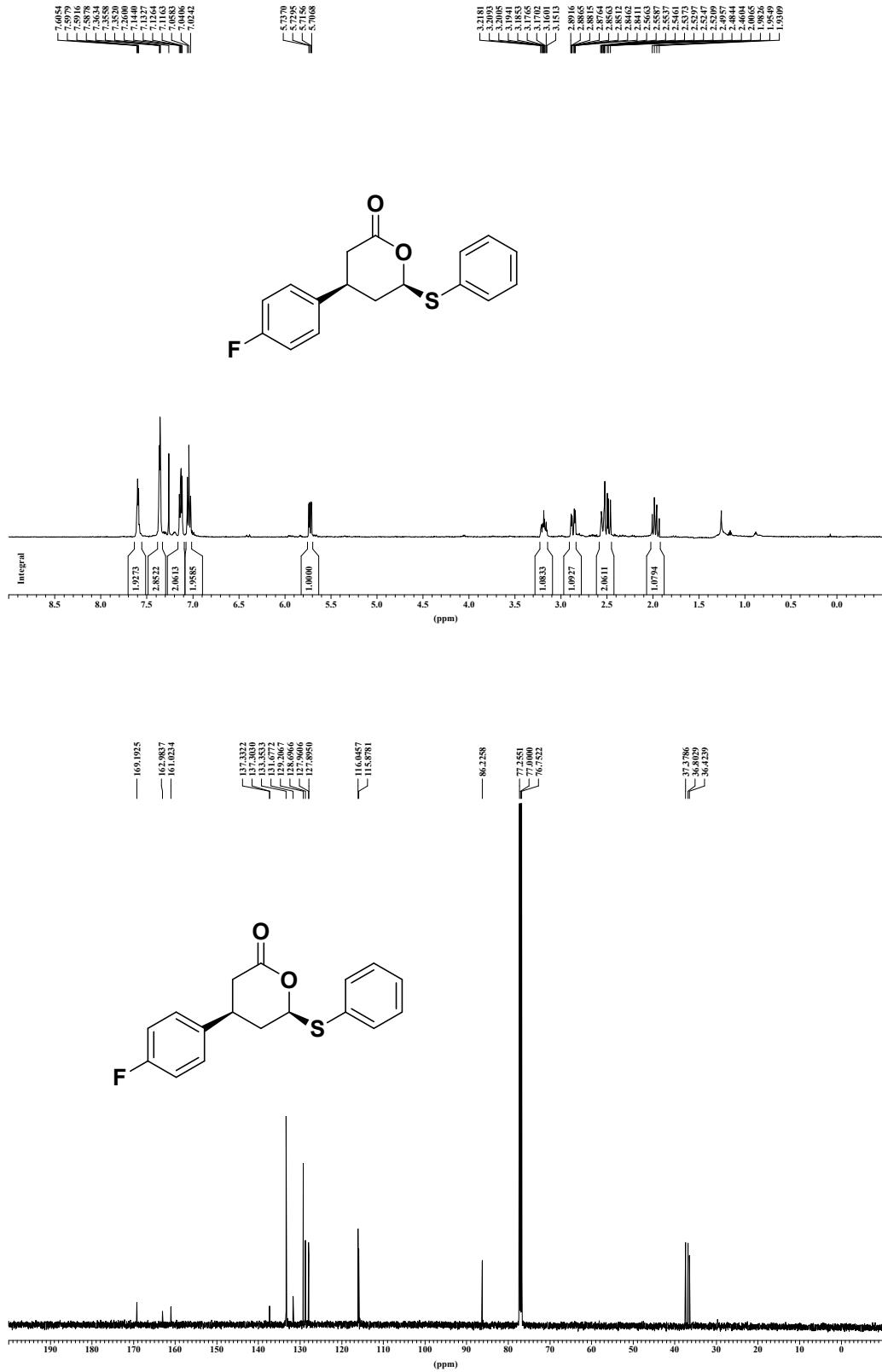
Compound d-2c



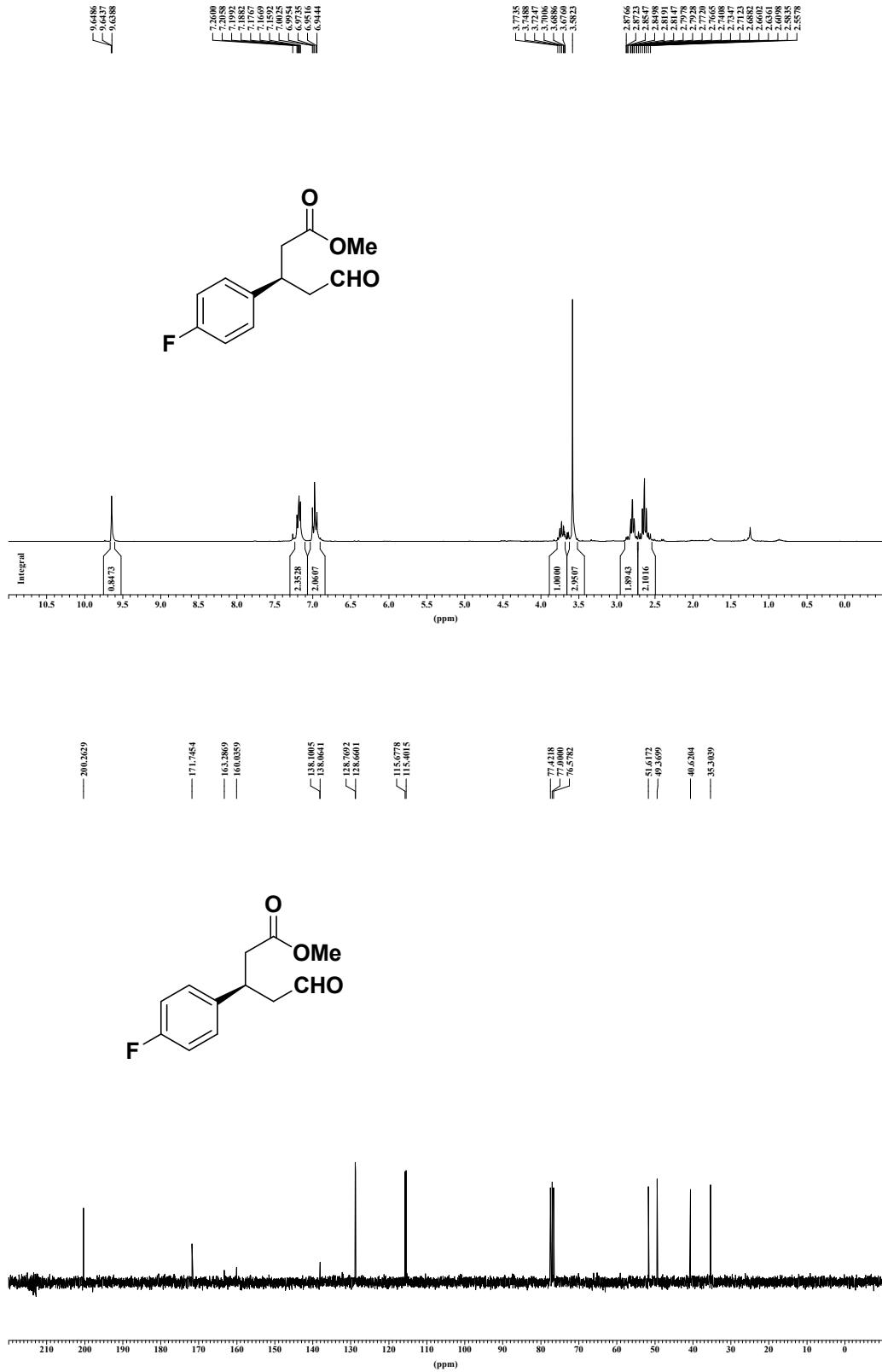
Compound *d*-3ac



Compound 3mc



Compound 5

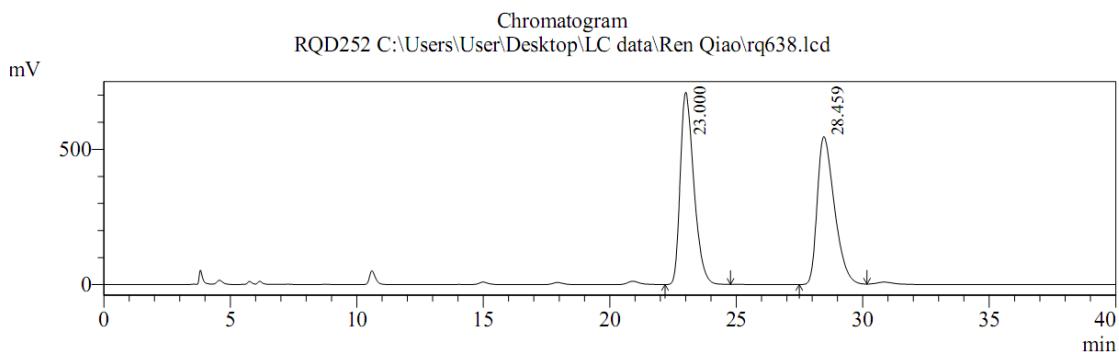
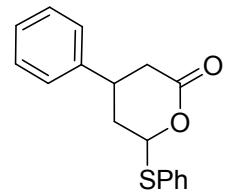


Compound 6



Racemic 3ac**===== Shimadzu LCsolution Analysis Report =====**

C:\Users\User\Desktop\LC data\Ren Qiao\rq638.lcd
 Acquired by : Admin
 Sample Name : RQD252
 Sample ID : RQ
 Data File Name : rq638.lcd
 Method File Name : 20%IPA, 1ml-min, 60min.lcm
 Batch File Name :
 Report File Name : Default.lcr
 Description : IC column ;20%IPA ;1ml/min

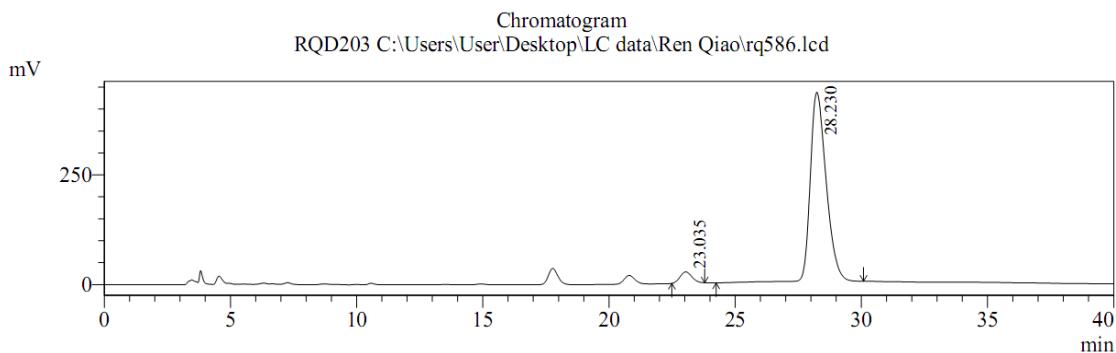
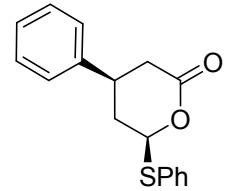


Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	23.000	26608734	710566	50.861	56.523
2	28.459	25708355	546556	49.139	43.477
Total		52317088	1257122	100.000	100.000

Enantiomeric enriched 3ac**===== Shimadzu LCsolution Analysis Report =====**

C:\Users\User\Desktop\LC data\Ren Qiao\rq586.lcd
 Acquired by : Admin
 Sample Name : RQD203
 Sample ID : RQ
 Data File Name : rq586.lcd
 Method File Name : 20%IPA, 1ml-min, 60min.lcm
 Batch File Name :
 Report File Name : Default.lcr
 Description : IC column ;20%IPA ;1ml/min



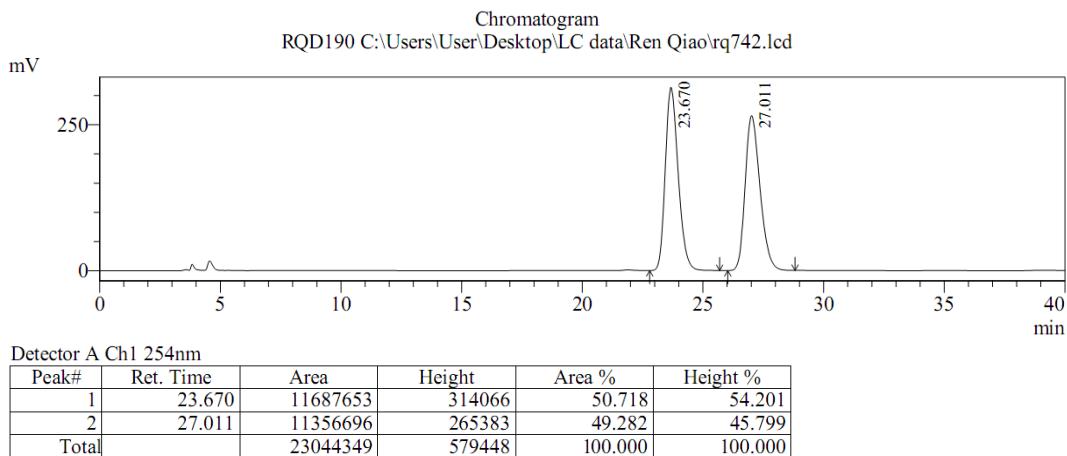
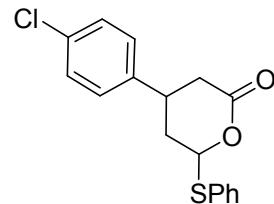
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	23.035	833667	25663	4.195	5.609
2	28.230	19038519	431887	95.805	94.391
Total		19872186	457551	100.000	100.000

Racemic 3bc

==== Shimadzu LCsolution Analysis Report ====

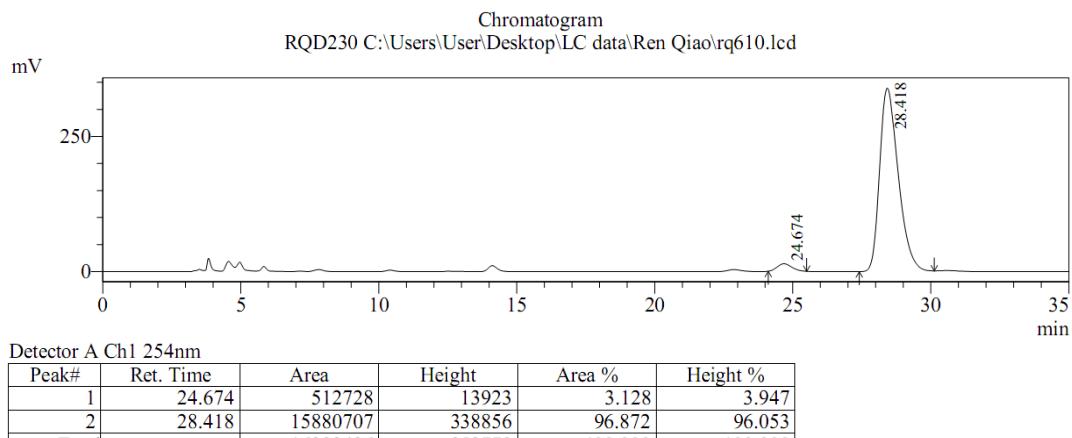
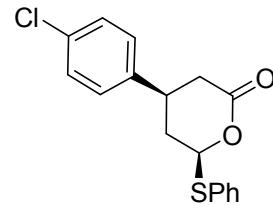
C:\Users\User\Desktop\LC data\Ren Qiao\rq742.lcd
Acquired by : Admin
Sample Name : RQD190
Sample ID : RQ
Data File Name : rq742.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column ;20%IPA ;1ml/min



Enantiomeric enriched 3bc

==== Shimadzu LCsolution Analysis Report ====

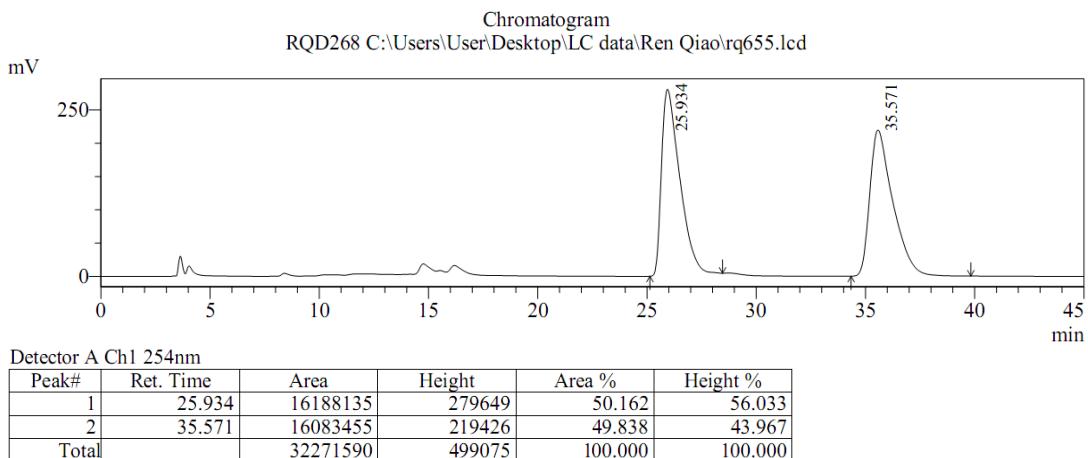
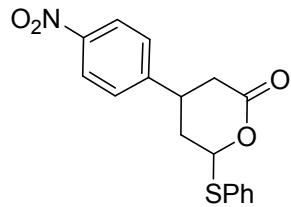
C:\Users\User\Desktop\LC data\Ren Qiao\rq610.lcd
Acquired by : Admin
Sample Name : RQD230
Sample ID : RQ
Data File Name : rq610.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column ;20%IPA ;1ml/min



Racemic 3cc

==== Shimadzu LCsolution Analysis Report ====

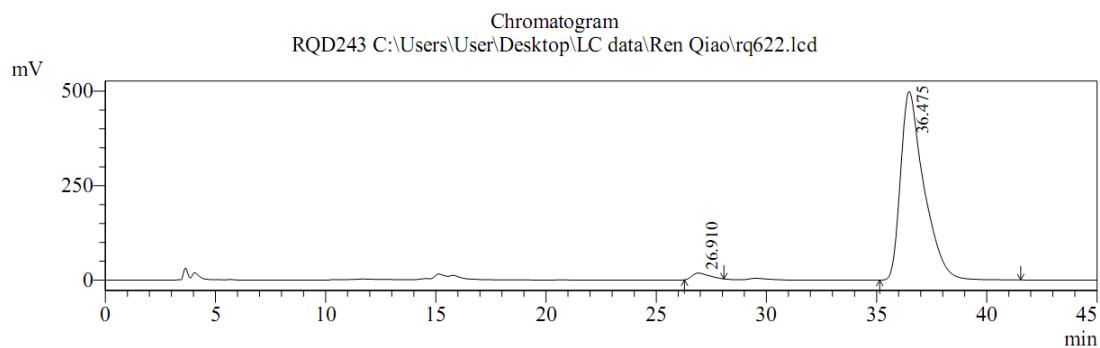
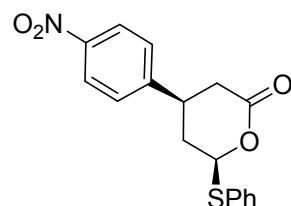
C:\Users\User\Desktop\LC data\Ren Qiao\rq655.lcd
Acquired by : Admin
Sample Name : RQD268
Sample ID : RQ
Data File Name : rq655.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IA column ;20%IPA ;1ml/min



Enantiomeric enriched 3cc

==== Shimadzu LCsolution Analysis Report ====

C:\Users\User\Desktop\LC data\Ren Qiao\rq622.lcd
Acquired by : Admin
Sample Name : RQD243
Sample ID : RQ
Data File Name : rq622.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IA column ;20%IPA ;1ml/min



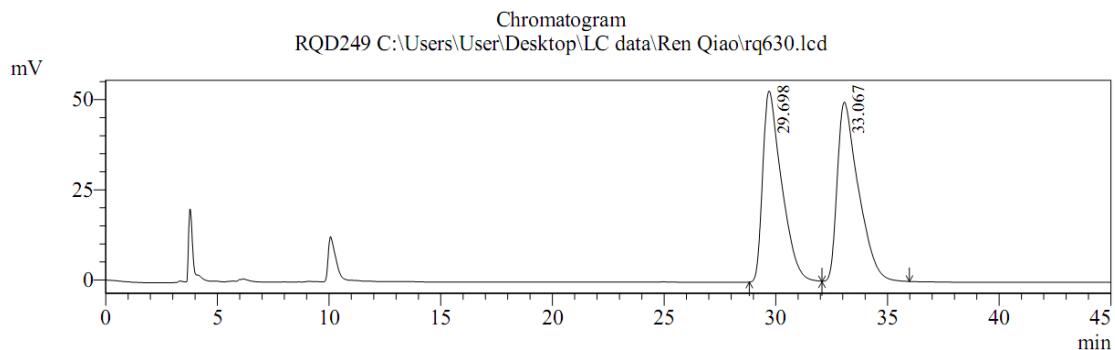
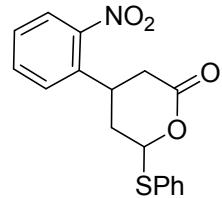
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	26.910	897338	17032	2.382	3.302
2	36.475	36767452	498801	97.618	96.698
Total		37664790	515834	100.000	100.000

Racemic 3dc

===== Shimadzu LCsolution Analysis Report =====

C:\Users\User\Desktop\LC data\Ren Qiao\rq630.lcd
Acquired by : Admin
Sample Name : RQD249
Sample ID : RQ
Data File Name : rq630.lcd
Method File Name : 10%IPA, 1ml-min, 40min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IA column ;10%IPA ;1ml/min



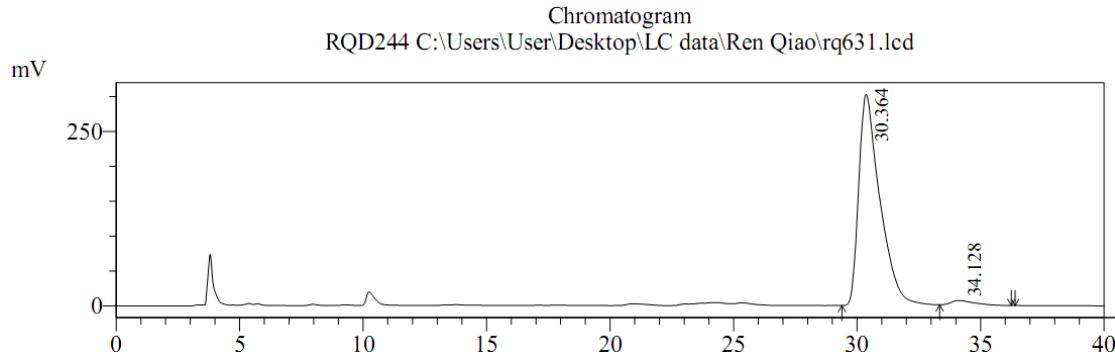
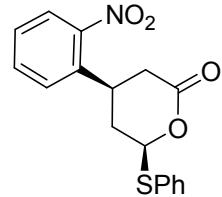
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	29.698	3127544	52949	48.780	51.536
2	33.067	3283933	49793	51.220	48.464
Total		6411476	102743	100.000	100.000

Enantiomeric enriched 3dc

===== Shimadzu LCsolution Analysis Report =====

C:\Users\User\Desktop\LC data\Ren Qiao\rq631.lcd
Acquired by : Admin
Sample Name : RQD244
Sample ID : RQ
Data File Name : rq631.lcd
Method File Name : 10%IPA, 1ml-min, 40min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IA column ;10%IPA ;1ml/min



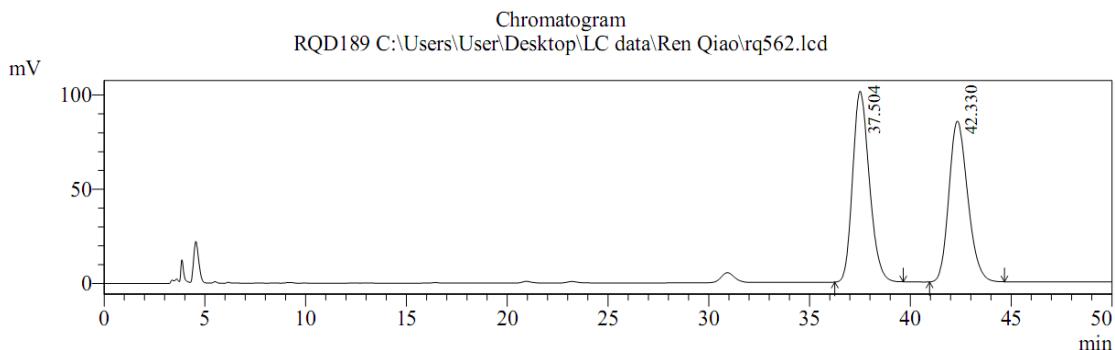
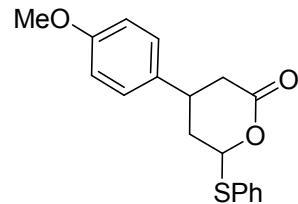
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	30.364	18182150	302361	97.820	97.935
2	34.128	405160	6375	2.180	2.065
Total		18587311	308735	100.000	100.000

Racemic 3ec

===== Shimadzu LCsolution Analysis Report =====

C:\Users\User\Desktop\LC data\Ren Qiao\rq562.lcd
Acquired by : Admin
Sample Name : RQD189
Sample ID : RQ
Data File Name : rq562.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column ;20%IPA ;1ml/min



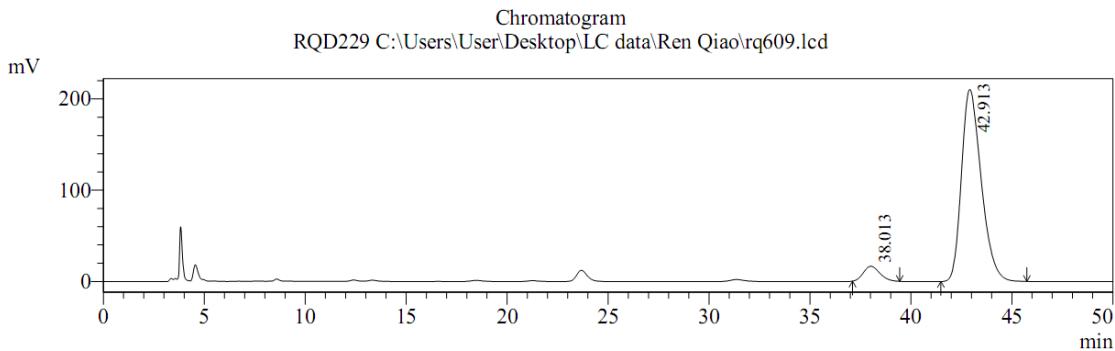
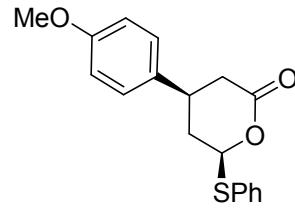
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	37.504	5838647	101256	51.328	54.254
2	42.330	5536554	85378	48.672	45.746
Total		11375202	186634	100.000	100.000

Enantiomeric enriched 3ec

===== Shimadzu LCsolution Analysis Report =====

C:\Users\User\Desktop\LC data\Ren Qiao\rq609.lcd
Acquired by : Admin
Sample Name : RQD229
Sample ID : RQ
Data File Name : rq609.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column ;20%IPA ;1ml/min

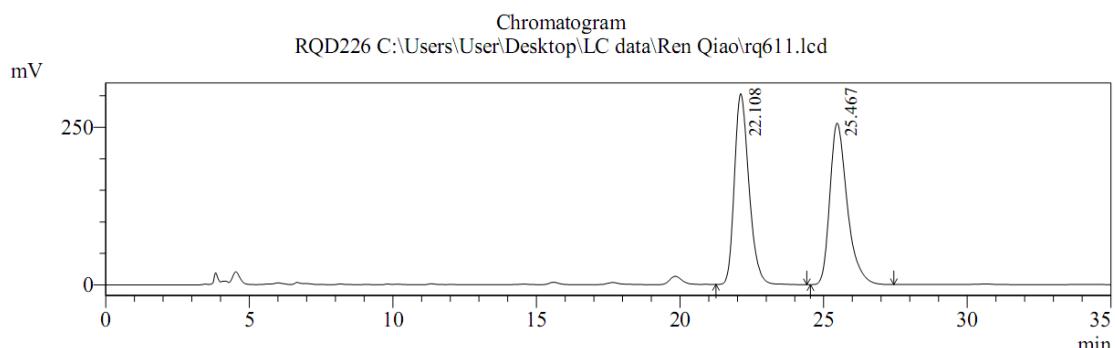
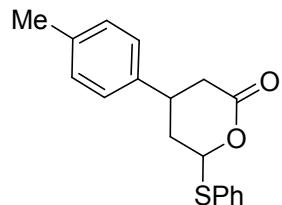


Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	38.013	926540	16419	6.075	7.233
2	42.913	14323915	210569	93.925	92.767
Total		15250455	226989	100.000	100.000

Racemic 3fc**==== Shimadzu LCsolution Analysis Report ====**

C:\Users\User\Desktop\LC data\Ren Qiao\rq611.lcd
 Acquired by : Admin
 Sample Name : RQD226
 Sample ID : RQ
 Data File Name : rq611.lcd
 Method File Name : 20%IPA, 1ml-min, 60min.lcm
 Batch File Name :
 Report File Name : Default.lcr
 Description : IC column ;20%IPA ;1ml/min

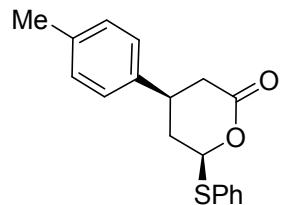


Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	22.108	10637781	302769	49.696	54.152
2	25.467	10767997	256339	50.304	45.848
Total		21405778	559109	100.000	100.000

Enantiomeric enriched 3fc**==== Shimadzu LCsolution Analysis Report ====**

C:\Users\User\Desktop\LC data\Ren Qiao\rq617.lcd
 Acquired by : Admin
 Sample Name : RQD233
 Sample ID : RQ
 Data File Name : rq617.lcd
 Method File Name : 20%IPA, 1ml-min, 60min.lcm
 Batch File Name :
 Report File Name : Default.lcr
 Description : IC column ;20%IPA ;1ml/min



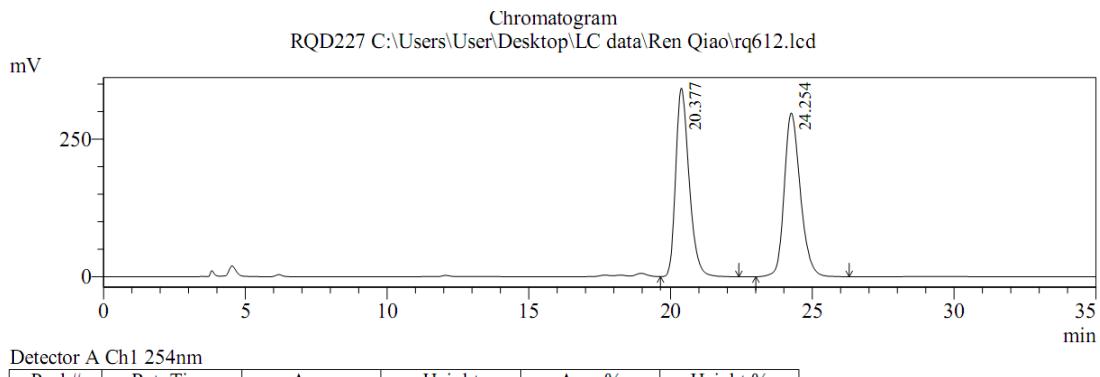
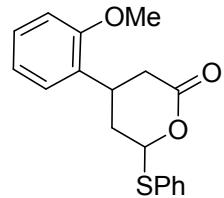
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	22.474	624113	20526	3.423	4.507
2	25.782	17609350	434892	96.577	95.493
Total		18233463	455418	100.000	100.000

Racemic 3gc

==== Shimadzu LCsolution Analysis Report ====

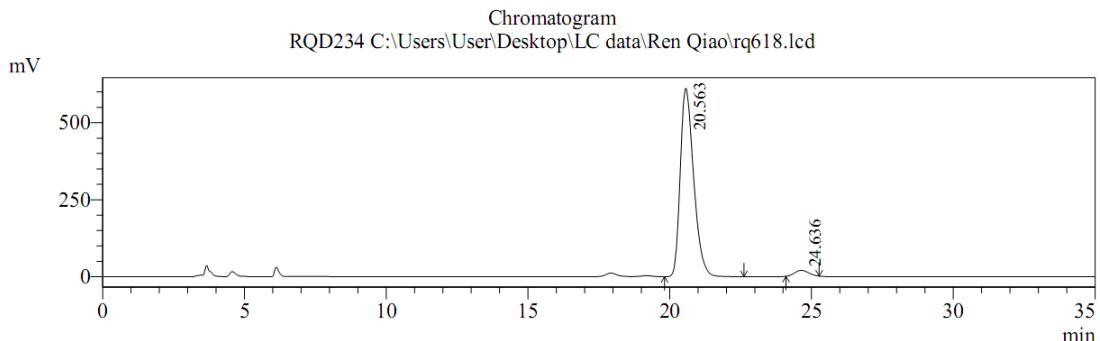
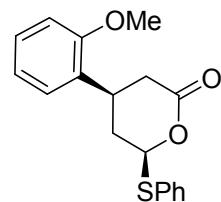
C:\Users\User\Desktop\LC data\Ren Qiao\rq612.lcd
Acquired by : Admin
Sample Name : RQD227
Sample ID : RQ
Data File Name : rq612.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column ;20%IPA ;1ml/min



Enantiomeric enriched 3gc

==== Shimadzu LCsolution Analysis Report ====

C:\Users\User\Desktop\LC data\Ren Qiao\rq618.lcd
Acquired by : Admin
Sample Name : RQD234
Sample ID : RQ
Data File Name : rq618.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column ;20%IPA ;1ml/min

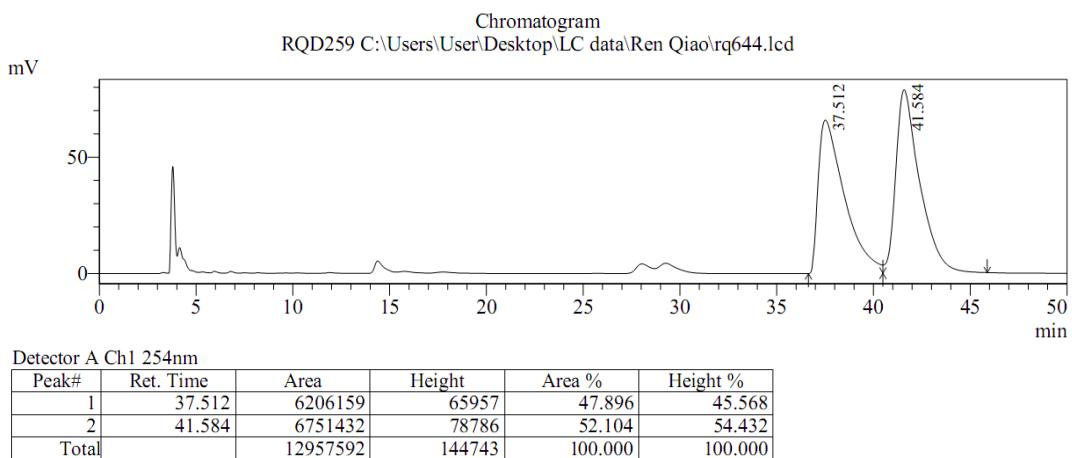
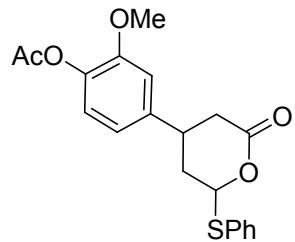


Peak#	Ret. Time	Area	Height	Area %	Height %
1	20.563	20067047	612236	96.845	96.949
2	24.636	653794	19268	3.155	3.051
Total		20720841	631504	100.000	100.000

Racemic 3hc

==== Shimadzu LCsolution Analysis Report ====

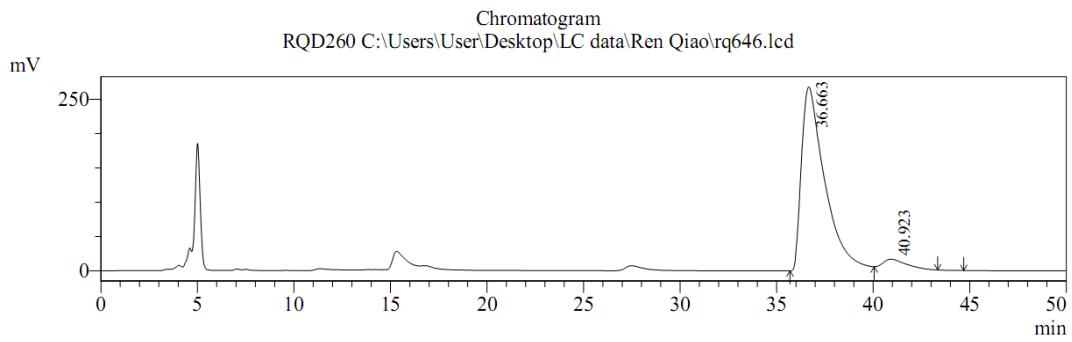
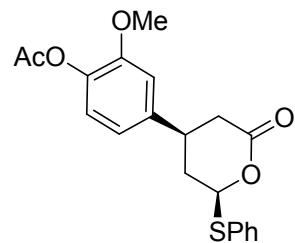
C:\Users\User\Desktop\LC data\Ren Qiao\rq644.lcd
 Acquired by : Admin
 Sample Name : RQD259
 Sample ID : RQ
 Data File Name : rq644.lcd
 Method File Name : 10%IPA, 1ml-min, 40min.lcm
 Batch File Name :
 Report File Name : Default.lcr
 Description : la column ;10%IPA ;1ml/min



Enantiomeric enriched 3hc

==== Shimadzu LCsolution Analysis Report ====

C:\Users\User\Desktop\LC data\Ren Qiao\rq646.lcd
 Acquired by : Admin
 Sample Name : RQD260
 Sample ID : RQ
 Data File Name : rq646.lcd
 Method File Name : 10%IPA, 1ml-min, 40min.lcm
 Batch File Name :
 Report File Name : Default.lcr
 Description : la column ;10%IPA ;1ml/min



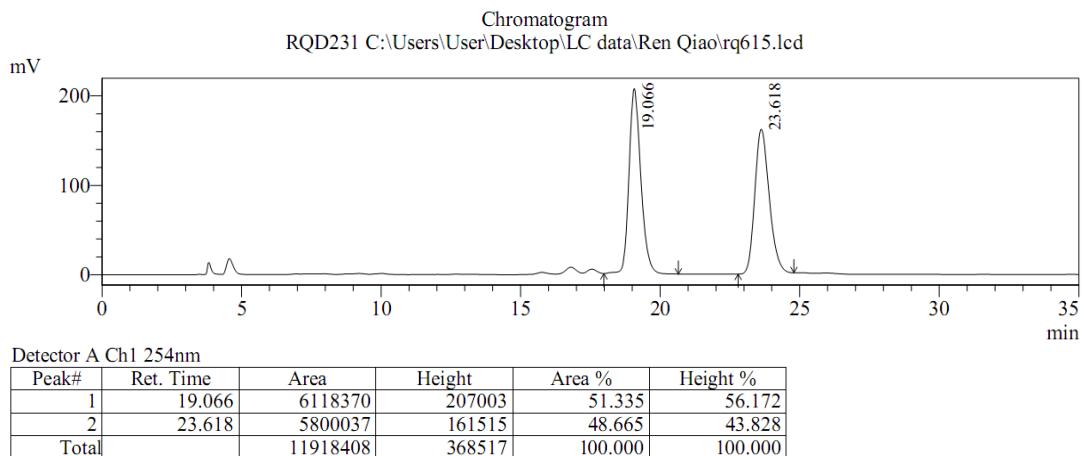
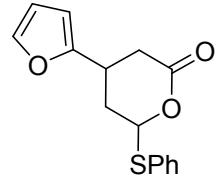
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	36.663	23362226	268178	96.062	95.617
2	40.923	957630	12292	3.938	4.383
Total		24319856	280470	100.000	100.000

Racemic 3ic

==== Shimadzu LCsolution Analysis Report ====

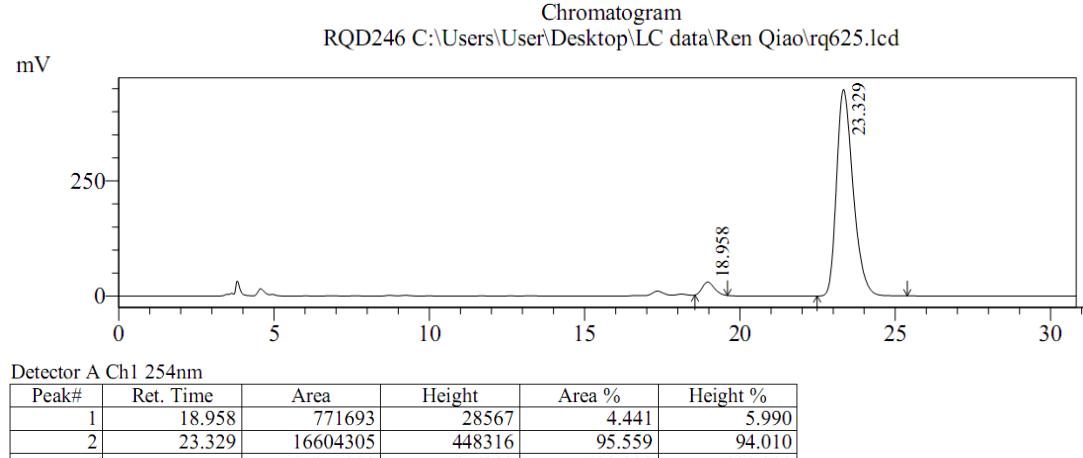
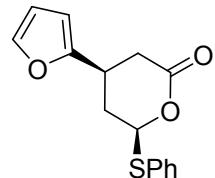
C:\Users\User\Desktop\LC data\Ren Qiao\rq615.lcd
Acquired by : Admin
Sample Name : RQD231
Sample ID : RQ
Data File Name : rq615.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column ;20%IPA ;1ml/min



Enantiomeric enriched 3ic

==== Shimadzu LCsolution Analysis Report ====

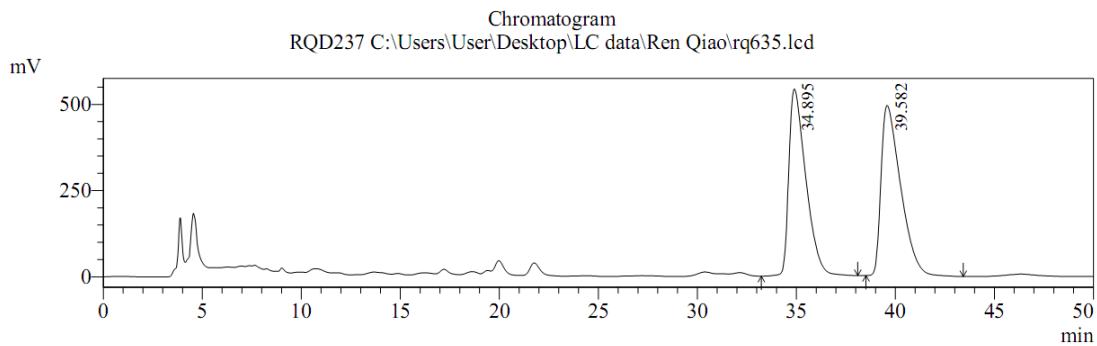
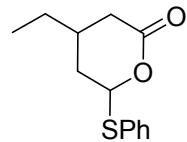
C:\Users\User\Desktop\LC data\Ren Qiao\rq625.lcd
Acquired by : Admin
Sample Name : RQD246
Sample ID : RQ
Data File Name : rq625.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column ;20%IPA ;1ml/min



Racemic 3jc

==== Shimadzu LCsolution Analysis Report ====

C:\Users\User\Desktop\LC data\Ren Qiao\rq635.lcd
Acquired by : Admin
Sample Name : RQD237
Sample ID : RQ
Data File Name : rq635.lcd
Method File Name : 10%IPA, 1ml-min, 40min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column ;10%IPA ;1ml/min



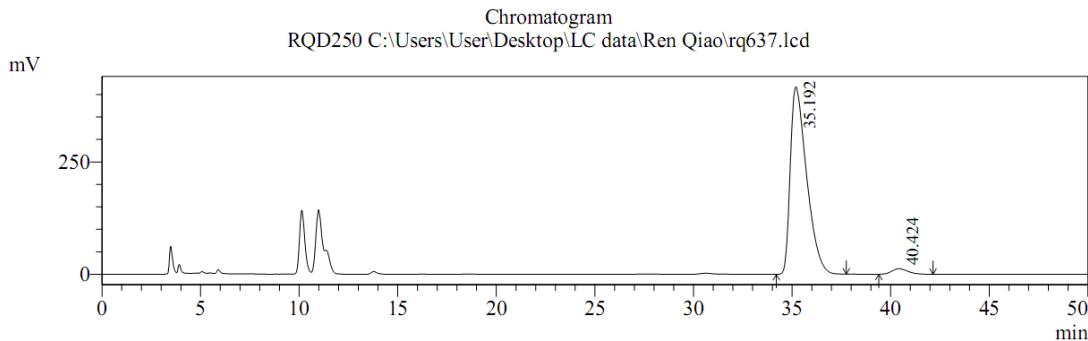
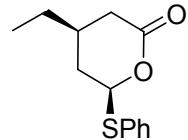
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	34.895	32378652	542495	48.831	52.346
2	39.582	33928905	493863	51.169	47.654
Total		66307557	1036357	100.000	100.000

Enantiomeric enriched 3jc

==== Shimadzu LCsolution Analysis Report ====

C:\Users\User\Desktop\LC data\Ren Qiao\rq637.lcd
Acquired by : Admin
Sample Name : RQD250
Sample ID : RQ
Data File Name : rq637.lcd
Method File Name : 10%IPA, 1ml-min, 40min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column ;10%IPA ;1ml/min



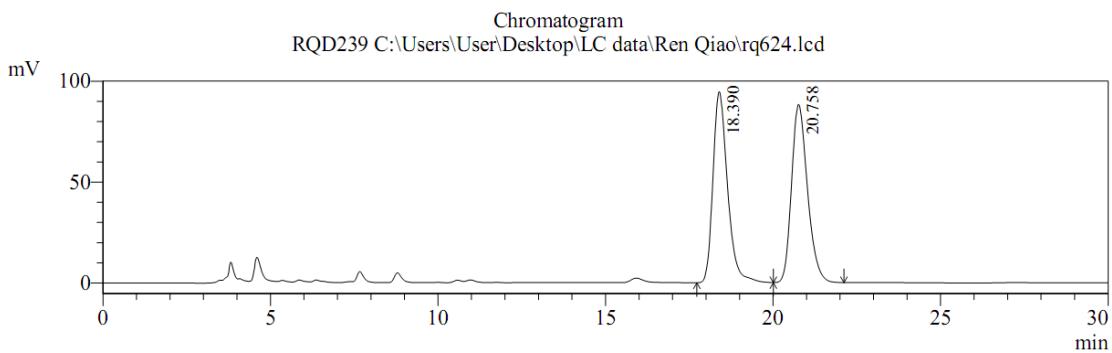
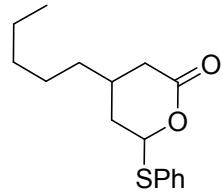
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	35.192	23985943	416814	97.144	97.099
2	40.424	705063	12453	2.856	2.901
Total		24691005	429267	100.000	100.000

Racemic 3kc

==== Shimadzu LCsolution Analysis Report ====

C:\Users\User\Desktop\LC data\Ren Qiao\rq624.lcd
Acquired by : Admin
Sample Name : RQD239
Sample ID : RQ
Data File Name : rq624.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column ;20%IPA ;1ml/min



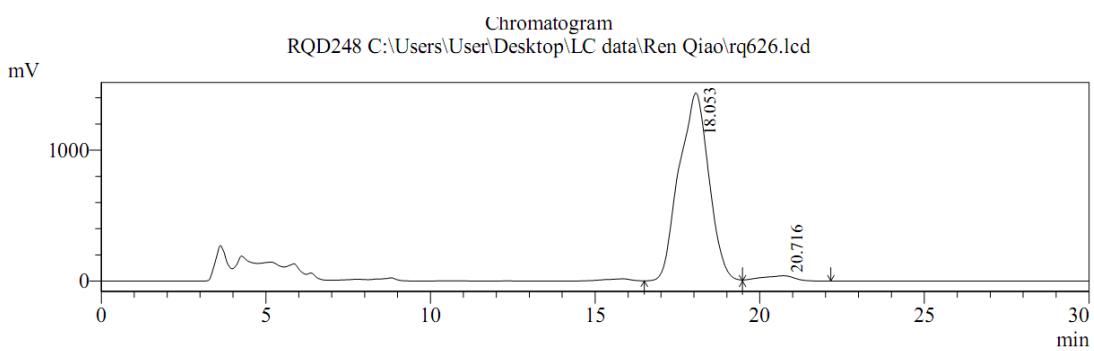
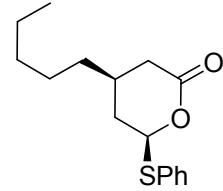
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	18.390	2859785	94585	49.460	51.736
2	20.758	2922266	88237	50.540	48.264
Total		5782051	182822	100.000	100.000

Enantiomeric enriched 3kc

==== Shimadzu LCsolution Analysis Report ====

C:\Users\User\Desktop\LC data\Ren Qiao\rq626.lcd
Acquired by : Admin
Sample Name : RQD248
Sample ID : RQ
Data File Name : rq626.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column ;20%IPA ;1ml/min



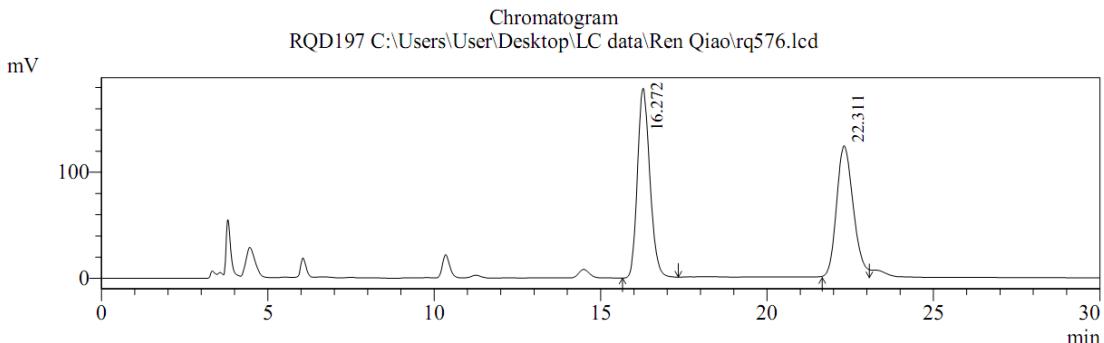
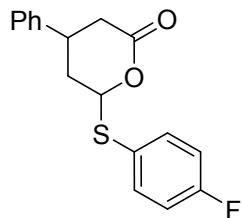
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	18.053	90445869	1435152	96.960	97.271
2	20.716	2835578	40270	3.040	2.729
Total		93281446	1475421	100.000	100.000

Racemic 3ad

==== Shimadzu LCsolution Analysis Report ====

C:\Users\User\Desktop\LC data\Ren Qiao\rq576.lcd
Acquired by : Admin
Sample Name : RQD197
Sample ID : RQ
Data File Name : rq576.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column ;20%IPA ;1ml/min



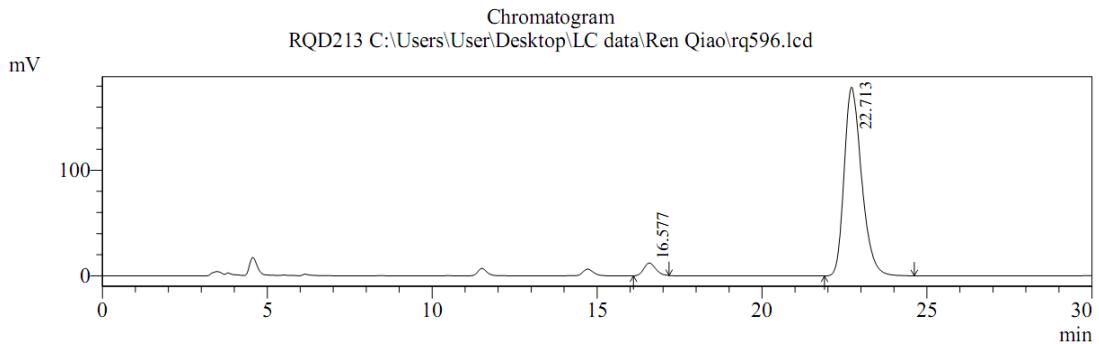
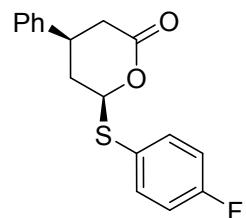
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	16.272	4630889	178695	51.427	58.966
2	22.311	4373877	124353	48.573	41.034
Total		9004766	303048	100.000	100.000

Enantiomeric enriched 3ad

==== Shimadzu LCsolution Analysis Report ====

C:\Users\User\Desktop\LC data\Ren Qiao\rq596.lcd
Acquired by : Admin
Sample Name : RQD213
Sample ID : RQ
Data File Name : rq596.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column ;20%IPA ;1ml/min



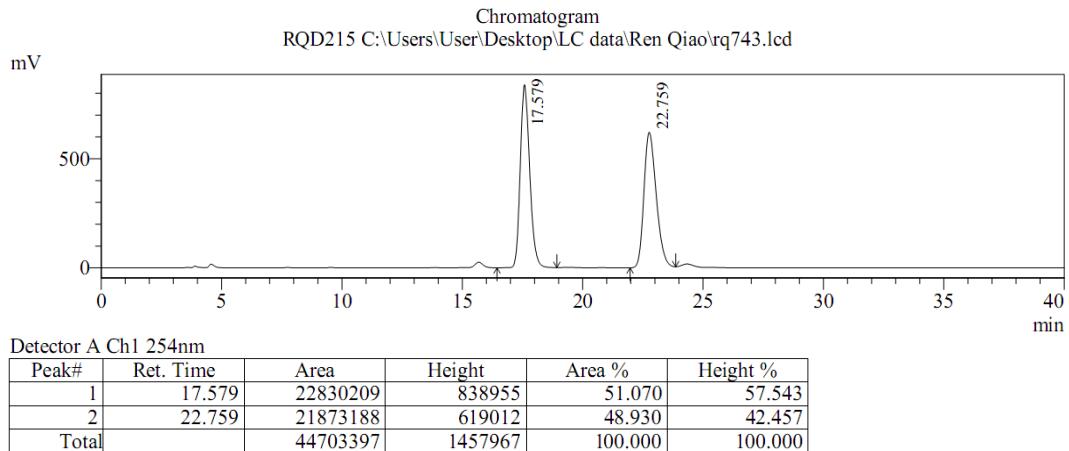
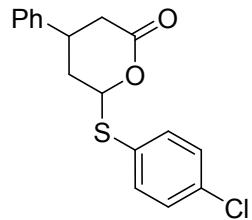
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	16.577	301209	12005	4.368	6.287
2	22.713	6594935	178957	95.632	93.713
Total		6896144	190963	100.000	100.000

Racemic 3ae

===== Shimadzu LCsolution Analysis Report =====

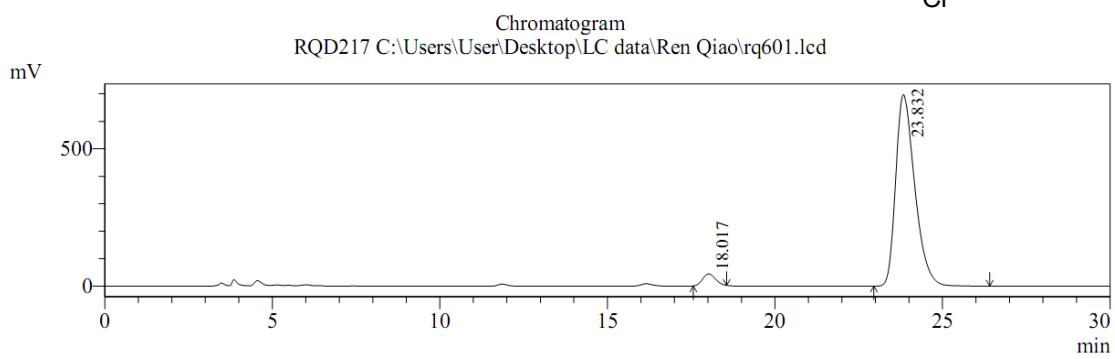
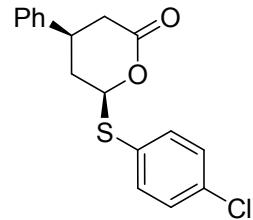
C:\Users\User\Desktop\LC data\Ren Qiao\rq743.lcd
 Acquired by : Admin
 Sample Name : RQD215
 Sample ID : RQ
 Data File Name : rq743.lcd
 Method File Name : 20%IPA, 1ml-min, 60min.lcm
 Batch File Name :
 Report File Name : Default.lcr
 Description : IC column ;20%IPA ;1ml/min



Enantiomeric enriched 3ae

===== Shimadzu LCsolution Analysis Report =====

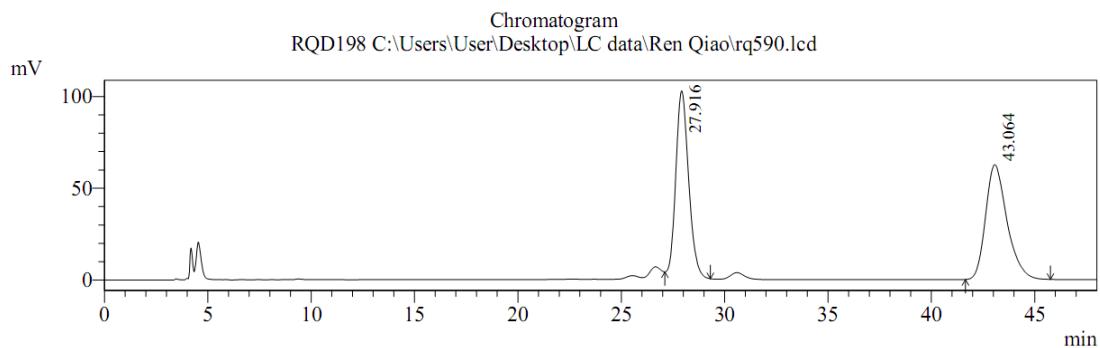
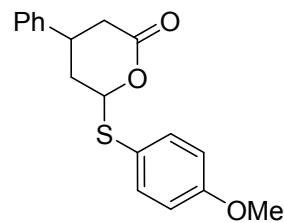
C:\Users\User\Desktop\LC data\Ren Qiao\rq601.lcd
 Acquired by : Admin
 Sample Name : RQD217
 Sample ID : RQ
 Data File Name : rq601.lcd
 Method File Name : 20%IPA, 1ml-min, 60min.lcm
 Batch File Name :
 Report File Name : Default.lcr
 Description : IC column ;20%IPA ;1ml/min



Peak#	Ret. Time	Area	Height	Area %	Height %
1	18.017	1135894	43070	3.957	5.818
2	23.832	27566808	697248	96.043	94.182
Total		28702702	740318	100.000	100.000

Racemic 3af**===== Shimadzu LCsolution Analysis Report =====**

C:\Users\User\Desktop\LC data\Ren Qiao\rq590.lcd
 Acquired by : Admin
 Sample Name : RQD198
 Sample ID : RQ
 Data File Name : rq590.lcd
 Method File Name : 20%IPA, 1ml-min, 60min.lcm
 Batch File Name :
 Report File Name : Default.lcr
 Description : IC column ;20%IPA ;1ml/min

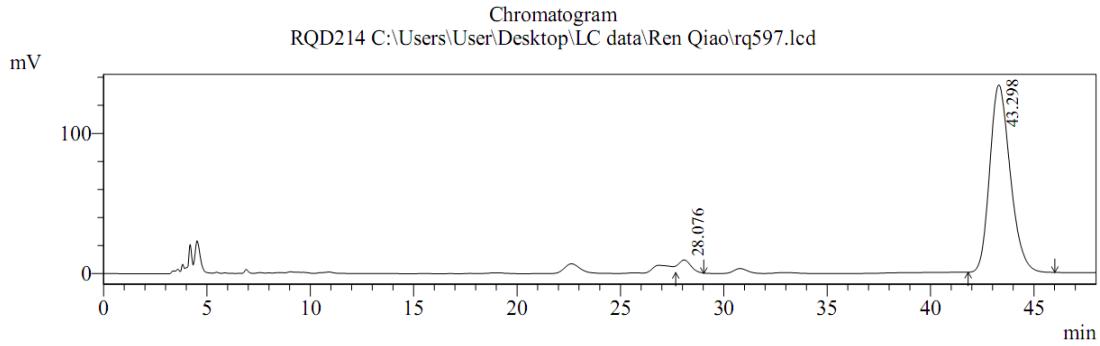
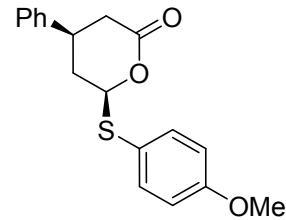


Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	27.916	4224521	99983	48.393	61.528
2	43.064	4505142	62517	51.607	38.472
Total		8729663	162500	100.000	100.000

Enantiomeric enriched 3af**===== Shimadzu LCsolution Analysis Report =====**

C:\Users\User\Desktop\LC data\Ren Qiao\rq597.lcd
 Acquired by : Admin
 Sample Name : RQD214
 Sample ID : RQ
 Data File Name : rq597.lcd
 Method File Name : 20%IPA, 1ml-min, 60min.lcm
 Batch File Name :
 Report File Name : Default.lcr
 Description : IC column ;20%IPA ;1ml/min

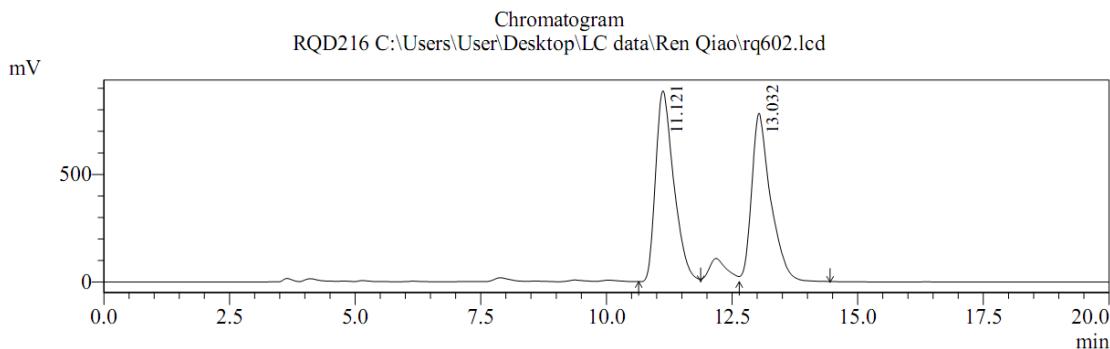
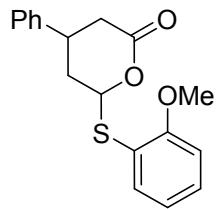


Detector A Ch1 254nm

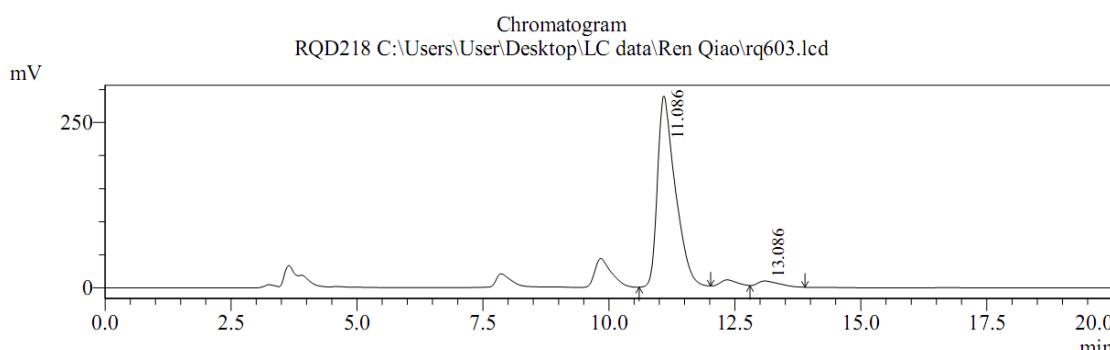
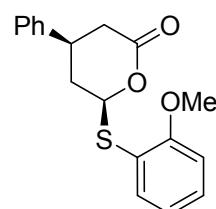
Peak#	Ret. Time	Area	Height	Area %	Height %
1	28.076	392577	9216	4.082	6.449
2	43.298	9225059	133679	95.918	93.551
Total		9617636	142895	100.000	100.000

Racemic 3ag**==== Shimadzu LCsolution Analysis Report ====**

C:\Users\User\Desktop\LC data\Ren Qiao\rq602.lcd
 Acquired by : Admin
 Sample Name : RQD216
 Sample ID : RQ
 Data File Name : rq602.lcd
 Method File Name : 20%IPA, 1ml-min, 60min.lcm
 Batch File Name :
 Report File Name : Default.lcr
 Description : IA column ;20%IPA ;1ml/min

**Enantiomeric enriched 3ag****==== Shimadzu LCsolution Analysis Report ====**

C:\Users\User\Desktop\LC data\Ren Qiao\rq603.lcd
 Acquired by : Admin
 Sample Name : RQD218
 Sample ID : RQ
 Data File Name : rq603.lcd
 Method File Name : 20%IPA, 1ml-min, 60min.lcm
 Batch File Name :
 Report File Name : Default.lcr
 Description : IA column ;20%IPA ;1ml/min



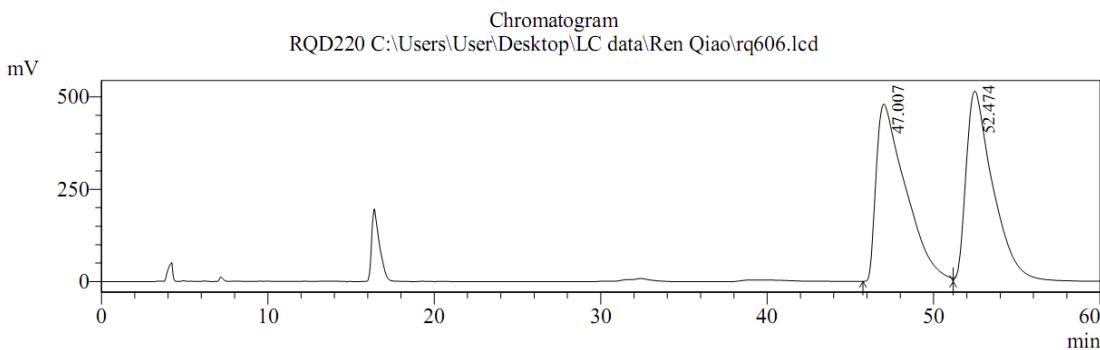
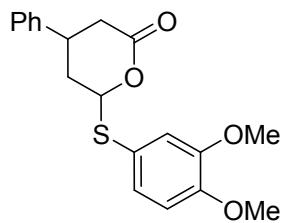
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	11.086	7098812	288673	97.104	97.392
2	13.086	211710	7731	2.896	2.608
Total		7310521	296404	100.000	100.000

Racemic 3ah

==== Shimadzu LCsolution Analysis Report ====

C:\Users\User\Desktop\LC data\Ren Qiao\rq606.lcd
Acquired by : Admin
Sample Name : RQD220
Sample ID : RQ
Data File Name : rq606.lcd
Method File Name : 5%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IA column ;5%IPA ;1ml/min



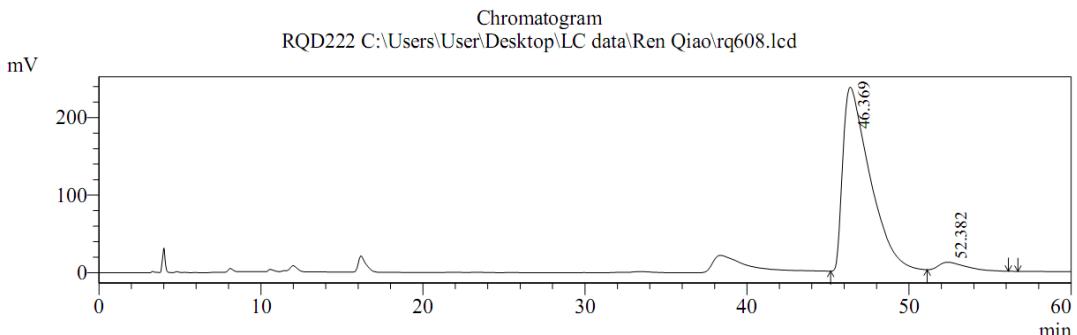
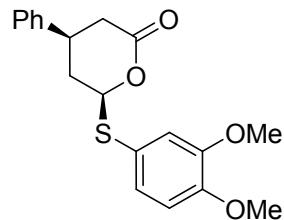
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	47.007	59266757	479863	50.548	48.225
2	52.474	57980817	515184	49.452	51.775
Total		117247574	995047	100.000	100.000

Enantiomeric enriched 3ah

==== Shimadzu LCsolution Analysis Report ====

C:\Users\User\Desktop\LC data\Ren Qiao\rq608.lcd
Acquired by : Admin
Sample Name : RQD222
Sample ID : RQ
Data File Name : rq608.lcd
Method File Name : 5%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IA column ;5%IPA ;1ml/min



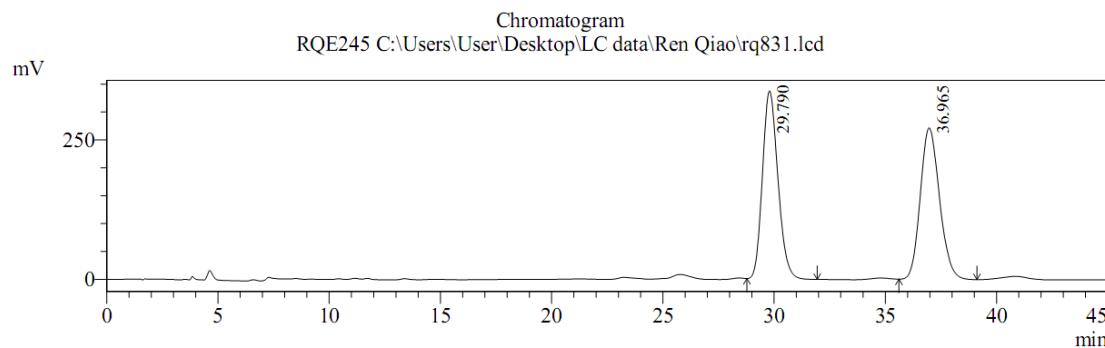
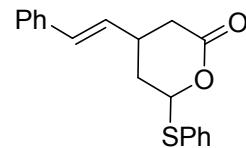
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	46.369	28065007	237134	96.206	95.909
2	52.382	1106878	10116	3.794	4.091
Total		29171885	247250	100.000	100.000

Racemic 3lc

==== Shimadzu LCsolution Analysis Report ====

C:\Users\User\Desktop\LC data\Ren Qiao\rq831.lcd
Acquired by : Admin
Sample Name : RQE245
Sample ID : RQ
Data File Name : rq831.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column with IC guard column ;20%IPA ;1 ml/min



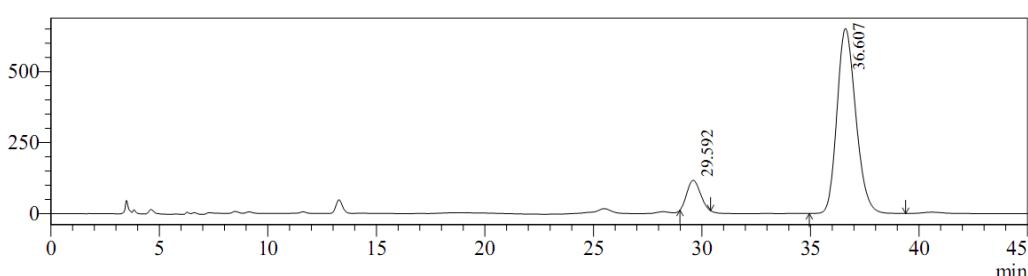
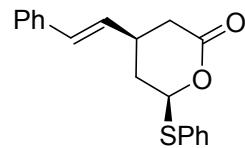
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	29.790	16257114	336926	49.968	55.364
2	36.965	16278012	271643	50.032	44.636
Total		32535126	608569	100.000	100.000

Enantiomeric enriched 3lc

==== Shimadzu LCsolution Analysis Report ====

C:\Users\User\Desktop\LC data\Ren Qiao\rq832.lcd
Acquired by : Admin
Sample Name : RQE247
Sample ID : RQ
Data File Name : rq832.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column with IC guard column ;20%IPA ;1 ml/min



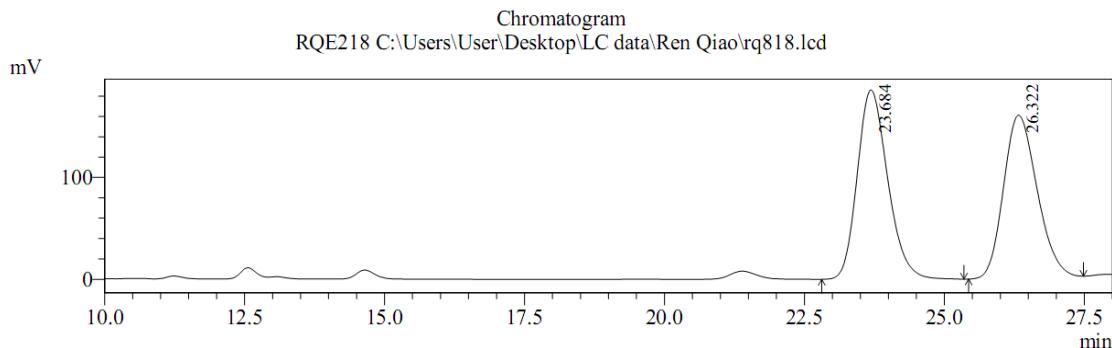
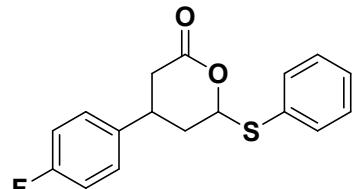
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	29.592	4457856	105749	10.216	13.976
2	36.607	39179395	650924	89.784	86.024
Total		43637250	756674	100.000	100.000

Racemic 3mc

==== Shimadzu LCsolution Analysis Report ====

C:\Users\User\Desktop\LC data\Ren Qiao\rq818.lcd
Acquired by : Admin
Sample Name : RQE218
Sample ID : RQ
Data File Name : rq818.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column with IC guard column ;20%IPA ;1.0 ml/min



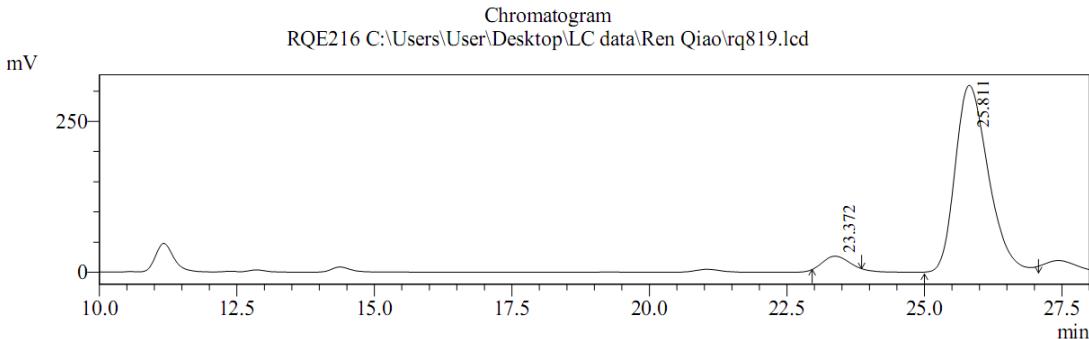
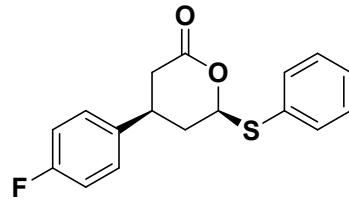
Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	23.684	7026566	185963	51.415	53.779
2	26.322	6639880	159827	48.585	46.221
Total		13666446	345790	100.000	100.000

Enantiomeric enriched 3mc

==== Shimadzu LCsolution Analysis Report ====

C:\Users\User\Desktop\LC data\Ren Qiao\rq819.lcd
Acquired by : Admin
Sample Name : RQE216
Sample ID : RQ
Data File Name : rq819.lcd
Method File Name : 20%IPA, 1ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description : IC column with IC guard column ;20%IPA ;1.0 ml/min



Detector A Ch1 254nm

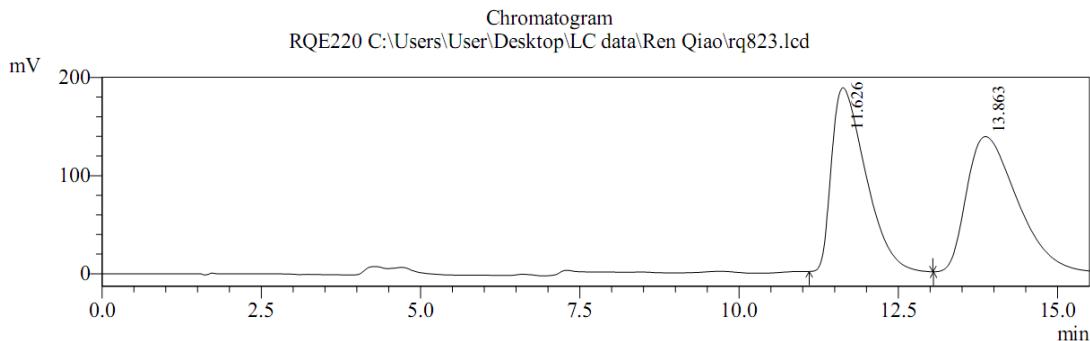
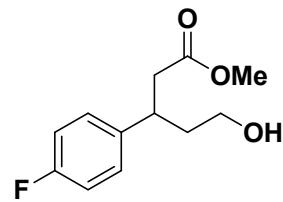
Peak#	Ret. Time	Area	Height	Area %	Height %
1	23.372	639316	21563	4.590	6.471
2	25.811	13290067	311654	95.410	93.529
Total		13929383	333216	100.000	100.000

Racemic 6

==== Shimadzu LCsolution Analysis Report ====

C:\Users\User\Desktop\LC data\Ren Qiao\rq823.lcd

Acquired by : Admin
Sample Name : RQE220
Sample ID : RQ
Data File Name : rq823.lcd
Method File Name : 10%IPA, 0.8ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description :OB column with IC guard column ;10%IPA ;0.8 ml/min



Detector A Ch1 254nm

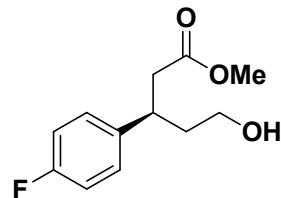
Peak#	Ret. Time	Area	Height	Area %	Height %
1	11.626	7002163	187405	48.057	57.625
2	13.863	7568298	137812	51.943	42.375
Total		14570461	325217	100.000	100.000

Enantiomeric enriched 18

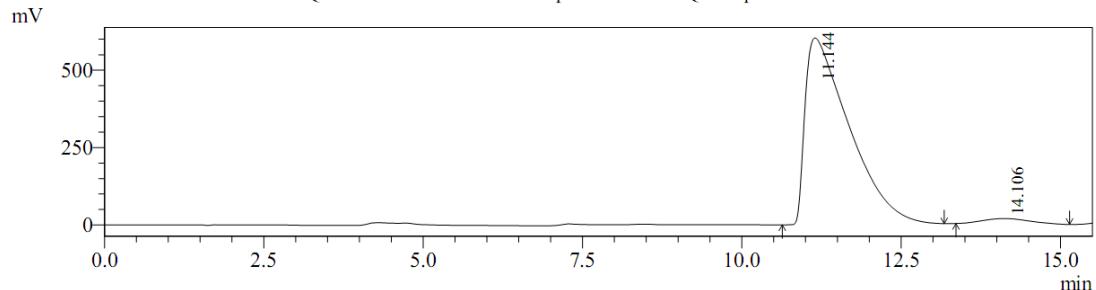
==== Shimadzu LCsolution Analysis Report ====

C:\Users\User\Desktop\LC data\Ren Qiao\rq824.lcd

Acquired by : Admin
Sample Name : RQE222
Sample ID : RQ
Data File Name : rq824.lcd
Method File Name : 10%IPA, 0.8ml-min, 60min.lcm
Batch File Name :
Report File Name : Default.lcr
Description :OB column with IC guard column ;10%IPA ;0.8 ml/min



Chromatogram
RQE222 C:\Users\User\Desktop\LC data\Ren Qiao\rq824.lcd



Detector A Ch1 254nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	11.144	28971570	602648	97.036	97.183
2	14.106	885070	17468	2.964	2.817
Total		29856640	620116	100.000	100.000