

Supporting Information for

Control of Four Stereocenters in an Organocatalytic Domino Double

Michael Reaction: Efficient Synthesis of Multisubstituted

Cyclopentanes

Bin Tan, Zugui Shi, Pei Juan Chua, and Guofu Zhong*

*Division of Chemistry and Biological Chemistry, School of Physical & Mathematical Sciences,
Nanyang Technological University, 21 Nanyang Link, Singapore 637371, Singapore*

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General Information: Analytical thin layer chromatography (TLC) was performed using Merck 60 F254 precoated silica gel plate (0.2 mm thickness). Subsequent to elution, plates were visualized using UV radiation (254 nm) on Spectroline Model ENF-24061/F 254 nm. Further visualization was possible by staining with basic solution of potassium permanganate or acidic solution of ceric molybdate.

Flash chromatography was performed using Merck silica gel 60 with freshly distilled solvents. Columns were typically packed as slurry and equilibrated with the appropriate solvent system prior to use.

Proton nuclear magnetic resonance spectra (^1H NMR) were recorded on Bruker AMX 400 spectrophotometer (CDCl_3 as solvent). Chemical shifts for ^1H NMR spectra are reported as δ in units of parts per million (ppm) downfield from SiMe_4 (δ 0.0) and relative to the signal of chloroform-d (δ 7.2600, singlet). Multiplicities were given as: s (singlet), d (doublet), t (triplet), dd (doublets of doublet) q (quartet) or m (multiplets). The number of protons (n) for a given resonance is indicated by nH. Coupling constants are reported as a *J* value in Hz. Carbon nuclear magnetic resonance spectra (^{13}C NMR) are reported as δ in units of parts per million (ppm) downfield

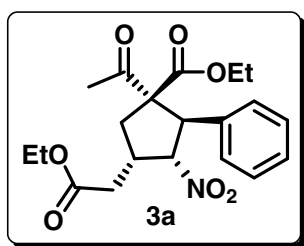
from SiMe₄ (δ 0.0) and relative to the signal of chloroform-d (δ 77.03, triplet).

Enantioselectivities were determined by High Performance Liquid Chromatography (HPLC) analysis (Shimadzu, LC-20AD) employing a Daicel Chirapak AD-H or AS-H column or Chiralcel OD-H. Optical rotations were measured in CH₂Cl₂ on a *Schmidt + Haensdch* polarimeter (Polartronic MH8) with a 10 cm cell (*c* given in g/100 mL).

High resolution mass spectrometry (HRMS) was recorded on Finnigan MAT 95 \times P spectrometer.

Typical Procedure for Double-Michael Reactions: To a solution of diethyl 5-acetylhex-2-enedioate (**1a**, 0.3 mmol, 1.0 eq) and nitro olefin (0.45 mmol, 1.5 eq) in diethyl ether (0.4 mL) was added catalyst **V** (**Q-NH₂**) (0.045 mmol, 0.15 eq) at room temperature (22°C). The resulting mixture was stirred vigorously for 16-36 hours. After the reaction was completed (monitored by TLC and crude NMR), the product was afforded by flash chromatography over silica gel (Et₂O:Hexane = 1:10 to 1:4).

(1*R*,2*R*,3*R*,4*S*)-ethyl 4-((ethoxycarbonyl)methyl)-1-acetyl-3-nitro-2-phenylcyclopentanecarboxylate (**3a**)



The title compound was prepared according to the typical procedure, as described above in 91% yield.

¹H-NMR (400 MHz, CDCl₃) δ 7.31-7.21 (m, 5H), 5.49 (dd, *J* = 5.2, 7.6 Hz, 1H), 5.10 (d, *J* = 4.8 Hz, 1H), 4.26-4.14 (m, 2H), 3.80-3.72 (m, 1H), 3.68-3.58 (m, 1H), 3.41-3.33 (m, 1H), 2.88 (dd, *J* = 6.8, 12.8 Hz, 1H), 2.50 (dd, *J* = 7.2, 16.8 Hz, 1H), 2.40 (dd, *J* = 7.6, 16.8 Hz, 1H), 2.20 (s, 3H), 2.01 (dd, *J* = 10.8, 12.8 Hz, 1H), 1.30 (t, *J* = 7.2 Hz, 3H), 0.76 (t, *J* = 7.2 Hz, 3H).

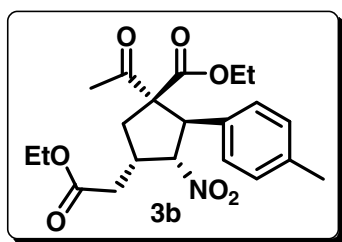
¹³C-NMR (100 MHz, CDCl₃) δ 200.09, 170.87, 170.29, 137.16, 128.52, 128.48, 127.80, 94.26, 71.28, 61.77, 61.05, 51.54, 40.27, 37.74, 34.20, 26.98, 14.16, 13.26.

HPLC: Chiralcel OD-H (hexane / *i*-PrOH = 80 / 20, flow rate 1 mL / min, λ = 230 nm), *t*_R (major) = 6.2 min, *t*_R (minor) = 8.9 min; 97% ee.

[α]_D²² = - 3.1 (*c* = 1.0, CH₂Cl₂).

HRMS (ESI) calcd for $C_{20}H_{26}NO_6+H$, m/z 392.1709, found 392.1707.

(1*R*,2*R*,3*R*,4*S*)-ethyl 4-((ethoxycarbonyl)methyl)-1-acetyl-3-nitro-2-*p*-tolylcyclopentanecarboxylate (**3b**)



The title compound was prepared according to the typical procedure, as described above in 89% yield.

1H -NMR (400 MHz, $CDCl_3$) δ 7.09 (m, 4H), 5.44 (dd, $J=5.2, 7.6$ Hz, 1H), 5.04 (d, $J = 5.2$ Hz, 1H), 4.24-4.15 (m, 2H), 3.80-3.75 (m, 1H), 3.63-3.57 (m, 1H), 3.45-3.41 (m, 1H), 2.87 (dd, $J = 6.8, 12.8$ Hz, 1H), 2.46 (dd, $J = 7.2, 16.8$ Hz, 1H), 2.40 (dd, $J = 7.6, 17.2$ Hz, 1H), 2.30 (s, 3H), 2.19 (s, 3H), 2.00 (dd, $J = 10.8, 12.8$ Hz, 1H), 1.29 (t, $J = 7.2$ Hz, 3H), 0.78 (t, $J = 7.2$ Hz, 3H).

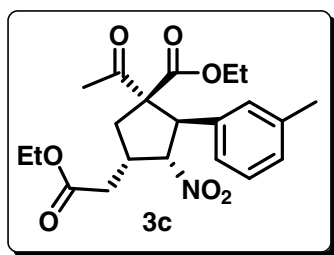
^{13}C -NMR (100 MHz, $CDCl_3$) δ 200.25, 170.89, 170.12, 137.52, 133.97, 129.12, 128.32, 94.28, 71.13, 61.74, 61.02, 51.29, 40.04, 37.77, 34.26, 27.04, 20.99, 14.15, 13.25.

HPLC: Chiralcel OD-H (hexane / *i*-PrOH = 80 / 20, flow rate 1 mL / min, λ = 220 nm), t_R (major) = 6.1 min, t_R (minor) = 8.7 min; 96% ee.

$[\alpha]_D^{22} = -5.7$ ($c = 1.3$, CH_2Cl_2).

HRMS (ESI) calcd for $\text{C}_{21}\text{H}_{27}\text{NO}_7 + \text{H}$, m/z 406.1866, found 406.1867.

(1*R*,2*R*,3*R*,4*S*)-ethyl 4-((ethoxycarbonyl)methyl)-1-acetyl-3-nitro-2-*m*-tolylcyclopentanecarboxylate (**3c**)



The title compound was prepared according to the typical procedure, as described above in 85% yield.

^1H -NMR (400 MHz, CDCl_3) δ 7.20-7.17 (m, 1H), 7.10-6.97 (m, 3H), 5.46 (m, 1H), 5.05 (d, $J = 5.2$ Hz, 1H), 4.24-4.16 (m, 2H), 3.80-3.72 (m, 1H), 3.64-3.58 (m, 1H), 3.45-3.36 (m, 1H), 2.87 (dd, $J = 6.8, 12.8$ Hz, 1H), 2.48 (dd, $J = 6.4, 17.2$ Hz, 1H), 2.36 (dd, $J = 7.6, 16.8$ Hz, 1H), 2.32 (s, 3H), 2.19 (s, 3H), 2.00 (dd, $J = 10.8, 12.8$ Hz, 1H), 1.30 (t, $J = 7.2$ Hz, 3H), 0.77 (t, $J = 7.2$ Hz, 3H).

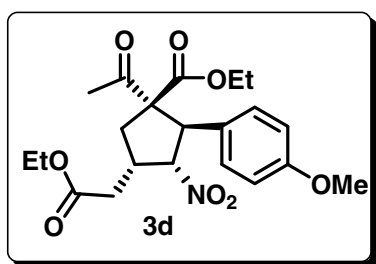
^{13}C -NMR (100 MHz, CDCl_3) δ 200.13, 170.88, 170.31, 138.13, 137.06, 129.33, 128.48, 128.41, 125.29, 94.36, 71.24, 61.71, 61.02, 51.53, 40.21, 37.77, 34.23, 26.99, 21.32, 14.16, 13.23.

HPLC: Chiralpak AD-H (hexane / *i*-PrOH = 90 / 10, flow rate 1 mL / min, $\lambda = 210$ nm), t_R (minor) = 10.6 min, t_R (major) = 12.0 min; 94% ee.

$[\alpha]_D^{22} = -4.2$ ($c = 1.2$, CH_2Cl_2).

HRMS (ESI) calcd for $C_{21}H_{27}NO_7+H$, m/z 406.1866, found 406.1858.

(1*R*,2*R*,3*R*,4*S*)-ethyl 4-((ethoxycarbonyl)methyl)-1-acetyl-2-(4-methoxyphenyl)-3-nitro-cyclopentanecarboxylate (**3d**)



To a solution of diethyl 5-acetylhex-2-enedioate (**1a**, 0.3 mmol, 1.0 eq) and 1-methoxy-4-((*E*)-2-nitrovinyl)benzene (0.6 mmol, 2.0 eq) in diethyl ether (0.4 mL) was added catalyst **V** (**Q-NH₂**) (0.06 mmol, 0.2 eq) at room temperature (22°C). The resulting mixture was stirred vigorously for 24 hours, then the reaction was continued for about 6 hours after removal of the solvent. After the reaction was completed (monitored by TLC and crude NMR), the title product was afforded by flash chromatography over silica gel (Et₂O:Hexane = 1:10 to 1:3) in 83% yield.

¹H-NMR (400 MHz, CDCl₃) δ 7.15 (d, J = 8.8 Hz, 2H), 6.82 (d, J = 8.8, Hz, 2H), 5.44 (dd, J = 5.6, 7.6 Hz, 1H), 5.02 (d, J = 5.2 Hz, 1H), 4.24-4.15 (m, 2H), 3.83-3.76 (m, 1H), 3.78 (s, 3H), 3.63-3.57 (m, 1H), 3.51-3.43 (m, 1H), 2.86 (dd, J = 6.8, 12.8 Hz, 1H), 2.48 (dd, J = 7.6, 16.8 Hz, 1H), 2.40 (dd, J = 7.6, 16.8 Hz, 1H), 2.19 (s, 3H), 2.00 (dd, J = 10.8, 12.8 Hz, 1H), 1.30 (t, J = 7.2 Hz, 3H), 0.83 (t, J = 7.2 Hz, 3H).

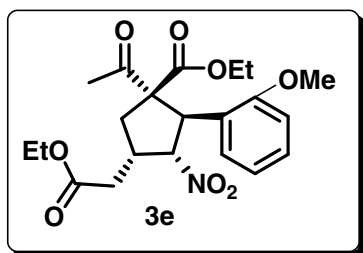
^{13}C -NMR (100 MHz, CDCl_3) δ 200.36, 170.90, 170.42, 159.14, 129.59, 128.96, 113.85, 94.32, 71.02, 61.77, 61.03, 55.31, 50.97, 39.88, 37.72, 34.27, 27.06, 14.15, 13.39.

HPLC: Chiralcel OD-H (hexane / *i*-PrOH = 80 / 20, flow rate 1 mL / min, λ = 220 nm), t_R (major) = 8.1 min, t_R (minor) = 12.2 min; 96% ee.

$[\alpha]_D^{22} = -9.0$ ($c = 1.3$, CH_2Cl_2).

HRMS (ESI) calcd for $\text{C}_{21}\text{H}_{27}\text{NO}_8 + \text{H}$, m/z 422.1815, found 422.1810.

(1*R*,2*R*,3*R*,4*S*)-ethyl 4-((ethoxycarbonylmethyl)-1-acetyl-2-(2-methoxyphenyl)-3-nitro-cyclopentanecarboxylate (**3e**)



To a solution of diethyl 5-acetylhex-2-enedioate (**1a**, 0.3 mmol, 1.0 eq) and 1-methoxy-2-((*E*)-2-nitrovinyl)benzene (0.6 mmol, 2.0 eq) in diethyl ether (0.4 mL) was added catalyst **V** (**Q-NH₂**) (0.06 mmol, 0.2 eq) at room temperature (22°C). The resulting mixture was stirred vigorously for 24 hours, then the reaction was continued for about 6 hours after removal of the solvent. After the reaction was completed (monitored by TLC and crude NMR), the title product was afforded by flash chromatography over silica gel (Et_2O :Hexane = 1:10 to 1:3) in 81% yield.

^1H -NMR (400 MHz, CDCl_3) δ 7.30-7.21 (m, 2H), 6.90-6.89 (m, 1H), 6.78 (d, $J = 8.0$ Hz, 1H), 5.55 (dd, $J = 4.0, 7.2$ Hz, 1H), 5.13 (d, $J = 4.0$ Hz, 1H), 4.21-4.13 (m, 2H), 3.90-3.84 (m, 1H), 3.77 (s, 3H), 3.65-3.59 (m, 1H), 3.53-3.45 (m, 1H), 2.89 (dd, $J = 6.4, 12.4$ Hz, 1H), 2.48 (dd, $J = 6.4, 17.2$ Hz, 1H), 2.34 (dd, $J = 8.4, 16.8$ Hz, 1H), 2.14 (s, 3H), 1.95 (m, 1H), 1.28 (t, $J = 7.2$ Hz, 3H), 0.78 (t, $J = 7.2$ Hz, 3H).

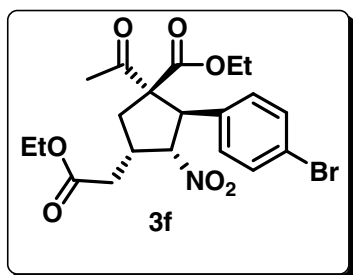
^{13}C -NMR (100 MHz, CDCl_3) δ 200.09, 171.03, 170.20, 157.31, 132.03, 129.12, 125.88, 120.85, 110.10, 94.46, 70.76, 61.39, 60.87, 54.90, 50.54, 39.78, 37.88, 34.10, 26.50, 14.19, 13.23.

HPLC: Chiralpak AD-H (hexane / *i*-PrOH = 90 / 10, flow rate 1 mL / min, $\lambda = 210$ nm), t_{R} (minor) = 12.4 min, t_{R} (major) = 14.6 min; 95% ee.

$[\alpha]_{\text{D}}^{22} = -5.5$ ($c = 1.0$, CH_2Cl_2).

HRMS (ESI) calcd for $\text{C}_{21}\text{H}_{27}\text{NO}_8 + \text{H}$, m/z 422.1815, found 422.1823.

(1*R*,2*R*,3*R*,4*S*)-ethyl 4-((ethoxycarbonyl)methyl)-1-acetyl-2-(4-bromophenyl)-3-nitro-cyclopentanecarboxylate (**3f**)



The title compound was prepared according to the typical procedure, as described above in 92% yield.

^1H -NMR (400 MHz, CDCl_3) δ 7.43 (d, $J = 8.4$ Hz, 1H), 7.11 (dd, $J = 8.4$ Hz, 1H), 5.44 (dd, $J = 5.6, 7.2$ Hz, 1H), 5.04 (d, $J = 5.2$ Hz, 1H), 4.24-4.16 (m, 2H), 3.86-3.78 (m, 1H), 3.63-3.57 (m, 1H), 3.53-3.45 (m, 1H), 2.86 (dd, $J = 6.8, 13.2$ Hz, 1H), 2.49 (dd, $J = 7.2, 16.8$ Hz, 1H), 2.39 (dd, $J = 7.6, 17.0$ Hz, 1H), 2.19 (s, 3H), 2.00 (dd, $J = 10.4, 12.4$ Hz, 1H), 1.30 (t, $J = 7.2$ Hz, 3H), 0.84 (t, $J = 7.2$ Hz, 3H).

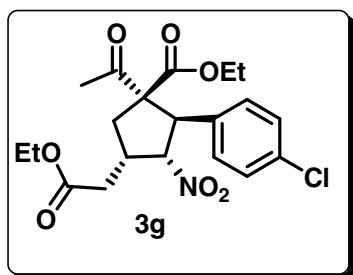
^{13}C -NMR (100 MHz, CDCl_3) δ 199.77, 170.81, 170.12, 136.02, 131.62, 130.21, 121.94, 93.72, 70.99, 61.96, 61.10, 50.93, 39.99, 37.70, 34.17, 26.96, 14.15, 13.33.

HPLC: Chiralcel OD-H (hexane / *i*-PrOH = 80 / 20, flow rate 1 mL / min, $\lambda = 220$ nm), t_{R} (major) = 7.6 min, t_{R} (minor) = 11.7 min; 97% ee.

$[\alpha]_{\text{D}}^{22} = -19.0$ ($c = 0.8$, CH_2Cl_2).

HRMS (ESI) calcd for $\text{C}_{20}\text{H}_{24}\text{BrNO}_7 + \text{H}$, m/z 470.0814, found 470.0808.

(1*R*,2*R*,3*R*,4*S*)-ethyl 4-((ethoxycarbonyl)methyl)-1-acetyl-2-(4-chlorophenyl)-3-nitro-cyclopentanecarboxylate (**3g**)



The title compound was prepared according to the typical procedure, as described above in 88% yield.

^1H -NMR (400 MHz, CDCl_3) δ 7.28 (d, $J = 6.8$ Hz, 1H), 7.16 (dd, $J = 2.0$, 7.8 Hz, 1H), 5.44 (dd, $J = 5.6$, 7.6 Hz, 1H), 5.05 (d, $J = 5.2$ Hz, 1H), 4.24-4.16 (m, 2H), 3.86-3.78 (m, 1H), 3.64-3.57 (m, 1H), 3.52-3.44 (m, 1H), 2.86 (dd, $J = 6.8$, 12.8 Hz, 1H), 2.48 (dd, $J = 7.4$, 17.0 Hz, 1H), 2.39 (dd, $J = 7.6$, 17.0 Hz, 1H), 2.19 (s, 3H), 2.00 (dd, $J = 10.4$, 12.4 Hz, 1H), 1.30 (t, $J = 7.2$ Hz, 3H), 0.84 (t, $J = 7.2$ Hz, 3H).

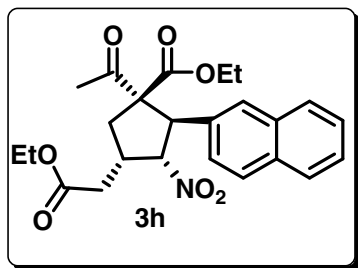
^{13}C -NMR (100 MHz, CDCl_3) δ 200.03, 170.81, 170.14, 135.49, 133.82, 129.88, 128.64, 93.79, 71.02, 61.92, 61.09, 50.86, 39.97, 37.70, 34.17, 26.97, 14.15, 13.33.

HPLC: Chiralcel OD-H (hexane / *i*-PrOH = 80 / 20, flow rate 1 mL / min, $\lambda = 254$ nm), t_{R} (major) = 7.1 min, t_{R} (minor) = 11.1 min; 96% ee.

$[\alpha]_{\text{D}}^{22} = -11.4$ ($c = 1.0$, CH_2Cl_2).

HRMS (ESI) calcd for $\text{C}_{20}\text{H}_{24}\text{ClNO}_7 + \text{H}$, m/z 426.1320, found 426.1322.

(1*R*,2*R*,3*R*,4*S*)-ethyl 4-((ethoxycarbonyl)methyl)-1-acetyl-2-(naphthalen-3-yl)-3-nitro-cyclopentanecarboxylate (**3h**)



The title compound was prepared according to the typical procedure, as described above in 84% yield.

$^1\text{H-NMR}$ (400 MHz, CDCl_3) δ 7.82-7.76 (m, 3H), 7.69 (s, 1H), 7.49-7.47(m, 2H), 7.34 (d, $J = 8.4$ Hz, 1H), 5.64-5.61 (m, 1H), 5.27 (d, $J = 5.2$ Hz, 1H), 4.26-4.18(m, 2H), 3.75-3.63 (m, 2H), 3.26-3.18 (m, 1H), 2.93 (dd, $J = 6.8, 12.8$ Hz, 1H), 2.53 (dd, $J = 7.6, 16.8$ Hz, 1H), 2.45 (dd, $J = 7.6, 16.8$ Hz, 1H), 2.21 (s, 3H), 2.45 (dd, $J = 10.8, 12.8$ Hz, 1H), 1.32 (t, $J = 7.2$ Hz, 3H), 0.52 (t, $J = 7.2$ Hz, 3H).

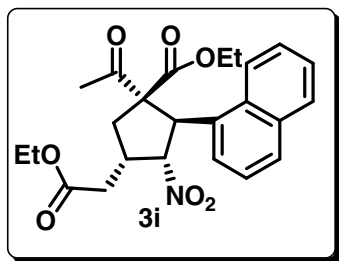
$^{13}\text{C-NMR}$ (100 MHz, CDCl_3) δ 200.22, 170.91, 170.32, 128.17, 127.89, 127.48, 127.43, 126.40, 126.28, 126.26, 94.23, 71/30, 61.74, 61.08, 51.71, 40.19, 37.86, 34.29, 27.02, 14.18, 13.03.

HPLC: Chiralpak AD-H (hexane / *i*-PrOH = 80 / 20, flow rate 1 mL / min, $\lambda = 254$ nm), t_{R} (minor) = 20.5 min, t_{R} (major) = 33.5 min; 95% ee.

$[\alpha]_{\text{D}}^{22} = -31.9$ ($c = 0.6$, CH_2Cl_2).

HRMS (ESI) calcd for $\text{C}_{24}\text{H}_{27}\text{NO}_7 + \text{H}$, m/z 442.1866, found 442.1861.

(1*R*,2*R*,3*R*,4*S*)-ethyl 4-((ethoxycarbonyl)methyl)-1-acetyl-2-(naphthalen-1-yl)-3-nitro-cyclopentanecarboxylate (**3i**)



The title compound was prepared according to the typical procedure, as described above in 87% yield.

$^1\text{H-NMR}$ (400 MHz, CDCl_3) δ 8.58 (d, $J = 7.6$ Hz, 1H), 7.82 (d, $J = 8.4$ Hz, 1H), 7.77 (d, $J = 8.4$ Hz, 1H), 7.62-7.59 (m, 1H), 7.52-7.49 (m, 1H), 7.44-7.40(m, 1H), 7.23 (d, $J = 7.2$ Hz, 1H), 6.03 (d, $J = 2.4$ Hz, 1H), 5.60-5.58 (m, 1H), 4.30-4.17(m, 2H), 3.76-3.68 (m, 1H), 3.51-3.45 (m, 1H), 3.01 (dd, $J = 6.8, 12.8$ Hz, 1H), 2.88-2.83 (m, 1H), 2.62 (dd, $J = 6.4, 17.2$ Hz, 1H), 2.43 (dd, $J = 8.4, 17.2$ Hz, 1H), 2.19 (s, 3H), 1.16-2.10 (m, 1H), 1.32 (t, $J = 7.2$ Hz, 3H), 0.19 (t, $J = 7.2$ Hz, 3H).

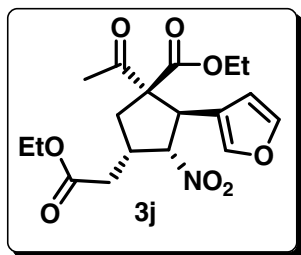
$^{13}\text{C-NMR}$ (100 MHz, CDCl_3) δ 199.62, 170.91, 170.19, 135.23, 133.55, 132.51, 128.57, 126.75, 126.04, 124.99, 124.75, 124.57, 96.98, 72.47, 61.48, 61.11, 46.43, 37.98, 33.91, 26.81, 14.19, 12.51.

HPLC: Chiralcel OD-H (hexane / *i*-PrOH = 80 / 20, flow rate 1 mL / min, $\lambda = 220$ nm), t_R (major) = 7.7 min, t_R (minor) = 10.2 min; 95% ee.

$[\alpha]_D^{22} = -29.0$ ($c = 1.0$, CH_2Cl_2).

HRMS (ESI) calcd for $\text{C}_{24}\text{H}_{27}\text{NO}_7 + \text{H}$, m/z 442.1866, found 442.1861.

(1*R*,2*R*,3*R*,4*S*)-ethyl 4-((ethoxycarbonyl)methyl)-1-acetyl-2-(furan-3-yl) 3-nitro-cyclopentanecarboxylate (**3j**)



The title compound was prepared according to the typical procedure, as described above in 86% yield.

$^1\text{H-NMR}$ (400 MHz, CDCl_3) δ 7.36-7.34 (m, 2H), 6.28 (s, 1H), 5.34-5.31(m, 1H), 4.82 (d, $J = 6.8$ Hz, 1H), 4.22-4.14(m, 2H), 4.03-3.95 (m, 1H), 3.88-3.81 (m, 1H), 3.58-3.52 (m, 1H), 2.83 (dd, $J = 7.2, 13.2$ Hz, 1H), 2.41 (d, $J = 7.6$ Hz, 1H), 2.21 (s, 3H), 1.96 (dd, $J = 10.4, 12.8$ Hz, 1H), 1.28 (t, $J = 7.2$ Hz, 3H), 1.02 (t, $J = 7.2$ Hz, 3H).

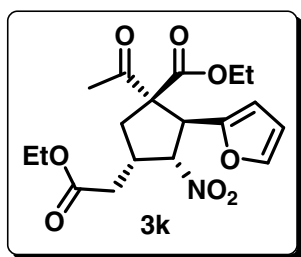
$^{13}\text{C-NMR}$ (100 MHz, CDCl_3) δ 200.61, 170.86, 170.57, 143.15, 140.79, 120.66, 109.98, 93.37, 69.47, 61.91, 61.04, 43.71, 38.29, 37.85, 34.55, 27.21, 14.14, 13.48.

HPLC: Chiralpak AD-H (hexane / *i*-PrOH = 90 / 10, flow rate 1 mL / min, $\lambda = 254$ nm), t_{R} (minor) = 17.9 min, t_{R} (major) = 19.0 min; 94% ee.

$[\alpha]_{\text{D}}^{22} = 5.7$ ($c = 1.2$, CH_2Cl_2).

HRMS (ESI) calcd for $\text{C}_{18}\text{H}_{23}\text{NO}_8 + \text{H}$, m/z 382.1502, found 382.1499.

(1*R*,2*R*,3*R*,4*S*)-ethyl 4-((ethoxycarbonyl)methyl)-1-acetyl-2-(furan-2-yl) 3-nitro-cyclopentanecarboxylate (**3k**)



The title compound was prepared according to the typical procedure, as described above in 87% yield.

$^1\text{H-NMR}$ (400 MHz, CDCl_3) δ 7.31-7.28 (m, 1H), 6.31-6.26 (m, 2H), 5.45 (dd, $J = 5.6, 7.6$ Hz, 1H), 5.09 (d, $J = 5.6$ Hz, 1H), 4.22-4.15(m, 2H),

4.04-3.99 (m, 1H), 3.80-3.75 (m, 1H), 3.60-3.53 (m, 1H), 2.88 (dd, $J = 6.8$, 13.2 Hz, 1H), 2.41 (d, $J = 7.6$ Hz, 1H), 2.20 (s, 3H), 1.96-1.90 (m, 1H), 1.28 (t, $J = 7.2$ Hz, 3H), 1.03 (t, $J = 7.2$ Hz, 3H).

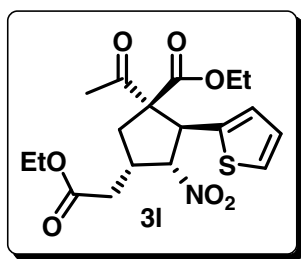
^{13}C -NMR (100 MHz, CDCl_3) δ 199.80, 170.76, 169.98, 149.81, 142.32, 110.73, 108.96, 92.07, 69.40, 62.32, 61.03, 46.32, 38.99, 37.50, 34.29, 26.73, 14.14, 13.60.

HPLC: Chiralcel OD-H (hexane / *i*-PrOH = 80 / 20, flow rate 1 mL / min, $\lambda = 220$ nm), t_R (major) = 7.2 min, t_R (minor) = 11.1 min; 94% ee.

$[\alpha]_D^{22} = -6.3$ ($c = 1.1$, CH_2Cl_2).

HRMS (ESI) calcd for $\text{C}_{18}\text{H}_{23}\text{NO}_8 + \text{Na}$, m/z 404.1321, found 404.1326.

(1*R*,2*R*,3*R*,4*S*) –ethyl 4-((ethoxycarbonyl)methyl)-1-acetyl-3-nitro-2-(thiophen-2-yl)cyclopentanecarboxylate (**3l**)



The title compound was prepared according to the typical procedure, as described above in 91% yield.

^1H -NMR (400 MHz, CDCl_3) δ 7.21-7.20 (m, 1H), 6.93-6.92 (m, 2H), 5.52-4.48 (m, 1H), 5.21 (d, $J = 7.2$ Hz, 1H), 4.226-4.14 (m, 2H), 3.96-3.89 (m, 1H), 3.75-3.68 (m, 1H), 3.66-3.57 (m, 1H), 2.89 (dd, $J = 7.2$, 12.8 Hz,

1H), 2.41 (d, $J = 7.6$ Hz, 1H), 2.21 (s, 3H), 1.96 (dd, $J = 10.0, 13.2$ Hz, 1H), 1.29 (t, $J = 7.2$ Hz, 3H), 0.94 (t, $J = 7.2$ Hz, 3H).

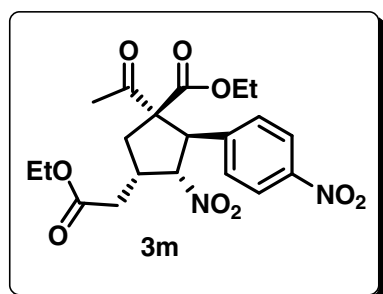
^{13}C -NMR (100 MHz, CDCl_3) δ 200.36, 170.76, 170.21, 138.64, 126.82, 126.27, 125.36, 94.23, 70.46, 62.10, 61.07, 47.35, 38.31, 37.70, 34.55, 27.14, 14.15, 13.48.

HPLC: Chiralpak AS-H (hexane / *i*-PrOH = 95 / 5, flow rate 1 mL / min, $\lambda = 210$ nm), t_{R} (minor) = 25.0 min, t_{R} (major) = 28.1 min; 94% ee.

$[\alpha]_{\text{D}}^{22} = -4.3$ ($c = 1.2$, CH_2Cl_2).

HRMS (ESI) calcd for $\text{C}_{18}\text{H}_{23}\text{NO}_7\text{S}+\text{Na}$, m/z 420.1093, found 420.1100.

(1*R*,2*R*,3*R*,4*S*)-ethyl 4-((ethoxycarbonyl)methyl)-1-acetyl-3-nitro-2-(4-nitrophenyl) cyclopentanecarboxylate (**3m**)



To a solution of diethyl 5-acetylhex-2-enedioate (**1a**, 0.3 mmol, 1.0 eq) and 1-nitro-4-((*E*)-2-nitrovinyl)benzene (0.6 mmol, 2.0 eq) in diethyl ether (0.4 mL) was added catalyst **V** (**Q-NH₂**) (0.06 mmol, 0.2 eq) at room temperature (22°C). The resulting mixture was stirred vigorously for 24 hours, then the reaction was continued for about 6 hours after removal of the solvent. After the reaction was completed (monitored by TLC and

crude NMR), the title product was afforded by flash chromatography over silica gel (Et₂O:Hexane = 1:10 to 1:3) in 81% yield.

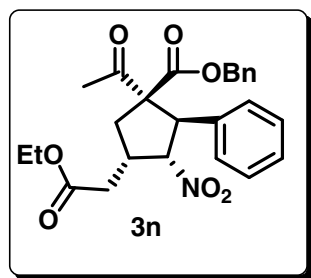
¹H-NMR (400 MHz, CDCl₃) δ 8.17 (d, *J* = 8.8 Hz, 1H), 7.44 (d, *J* = 8.8 Hz, 1H), 5.52-5.49 (m, 1H), 5.19 (d, *J* = 5.6 Hz, 1H), 4.25-4.16 (m, 2H), 3.86-3.80 (m, 1H), 3.69-3.62 (m, 1H), 3.49-3.43 (m, 1H), 2.89 (dd, *J* = 6.8, 13.2 Hz, 1H), 2.51 (dd, *J* = 7.2, 17.2 Hz, 1H), 2.40 (dd, *J* = 7.6, 17.2 Hz, 1H), 2.22 (s, 3H), 2.02 (dd, *J* = 10.8, 12.8 Hz, 1H), 1.30 (t, *J* = 7.2 Hz, 3H), 0.82 (t, *J* = 7.2 Hz, 3H).

¹³C-NMR (100 MHz, CDCl₃) δ 199.79, 170.73, 169.82, 147.41, 144.35, 129.64, 123.60, 93.20, 71.10, 62.07, 61.21, 51.02, 40.01, 37.73, 34.09, 26.91, 14.15, 13.42.

HPLC: Chiralpak AS-H (hexane / *i*-PrOH = 80 / 20, flow rate 1 mL / min, λ = 210 nm), *t*_R (major) = 20.9 min, *t*_R (minor) = 25.4 min; 97% ee.

[α]_D²² = - 9.7 (*c* = 1.0, CH₂Cl₂).

(1*R*,2*R*,3*R*,4*S*)-benzyl 4-((ethoxycarbonyl)methyl)-1-acetyl-3-nitro-2-phenylcyclopentanecarboxylate (**3n**)



To a solution of 6-benzyl 1-ethyl 5-acetylhex-2-enedioate **1b** (0.3 mmol, 1.0 eq) and 1-((*E*)-2-nitrovinyl)benzene (0.45 mmol, 1.5 eq) in diethyl

ether (0.4 mL) was added catalyst **V** (**Q-NH₂**) (0.045 mmol, 0.15 eq) at room temperature (22°C). The resulting mixture was stirred vigorously for 16 hours, then the reaction was continued for about 6 hours after removal of the solvent. After the reaction was completed (monitored by TLC and crude NMR), the title product was afforded by flash chromatography over silica gel (Ethyl acetate:Hexane = 1:10 to 1:4) in 91 % yield.

¹H-NMR (400 MHz, CDCl₃) δ 7.54-7.14 (m, 8H), 6.96-6.94 (m, 2H), 5.51 (dd, *J* = 5.2, 7.2 Hz, 1H), 5.11 (d, *J* = 5.2 Hz, 1H), 4.83 (d, *J* = 12.0 Hz, 1H), 4.25-4.16 (m, 2H), 4.10 (d, *J* = 12.0 Hz, 1H), 3.69-3.63 (m, 1H), 2.89 (dd, *J* = 6.8, 13.2 Hz, 1H), 2.49 (dd, *J* = 7.6, 16.8 Hz, 1H), 2.40 (dd, *J* = 7.6, 16.8 Hz, 1H), 2.11 (s, 3H), 2.02 (dd, *J* = 10.8, 12.8 Hz, 1H), 1.30 (t, *J* = 7.2 Hz, 3H).

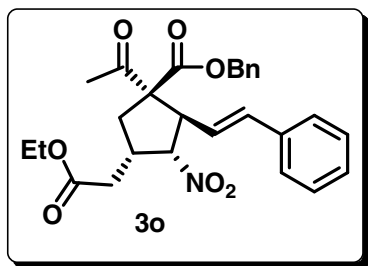
¹³C-NMR (100 MHz, CDCl₃) δ 199.84, 170.86, 170.18, 137.04, 134.09, 128.63, 128.49, 128.45, 128.28, 127.91, 94.23, 71.41, 67.57, 61.07, 51.69, 40.20, 37.81, 34.22, 27.07, 14.17.

HPLC: Chiralcel OD-H (hexane / *i*-PrOH = 80 / 20, flow rate 1 mL / min, λ = 220 nm), *t_R* (major) = 7.7 min, *t_R* (minor) = 11.5 min; 97% ee.

[α]_D²² = - 5.0 (*c* = 0.9, CH₂Cl₂).

HRMS (ESI) calcd for C₂₅H₂₇NO₇+H, *m/z* 454.1866, found 454.1861.

(1*R*,2*R*,3*R*,4*S*)-benzyl 4-((ethoxycarbonyl)methyl)-1-acetyl-3-nitro-2-styrylcyclopentanecarboxylate (**3o**)



To a solution of 6-benzyl 1-ethyl 5-acetylhex-2-enedioate (**1b**, 0.3 mmol, 1.0 eq) and 1-((1*E*,3*E*)-4-nitrobuta-1,3-dienyl)benzene (0.6 mmol, 2.0 eq) in diethyl ether (0.4 mL) was added catalyst **V** (**Q-NH₂**) (0.03 mmol, 0.2 eq) at room temperature (22°C). The resulting mixture was stirred vigorously for 24 hours, and then the reaction was continued for about 6 hours after removal of the solvent. After the reaction was completed (monitored by TLC and crude NMR), the title product was afforded by flash chromatography over silica gel (Et₂O:Hexane = 1:10 to 1:4) in 83% yield.

¹H-NMR (400 MHz, CDCl₃) δ 7.32-7.22 (m, 10H), 6.55 (d, *J* = 16.0, 1H), 6.05 (dd, *J* = 8.8, 16.0 Hz, 1H), 5.21-5.17 (m, 1H), 5.18 (d, *J* = 12.0 Hz, 1H), 5.05 (d, *J* = 12.0 Hz, 1H), 4.08 (d, *J* = 12.4 Hz, 1H), 4.26-4.13 (m, 3H), 3.45-3.35 (m, 1H), 2.72 (dd, *J* = 7.2, 13.2 Hz, 1H), 2.43-2.40 (m, 2H), 2.18 (s, 3H), 2.07 (dd, *J* = 10.8, 13.2 Hz, 1H), 1.28 (t, *J* = 7.2 Hz, 3H).

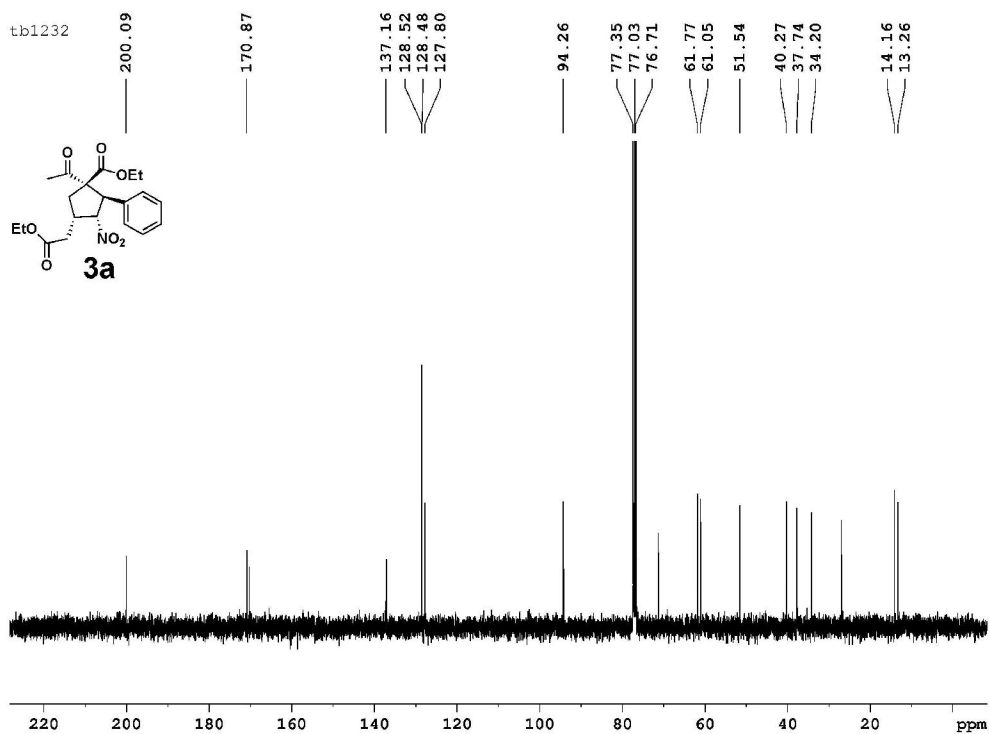
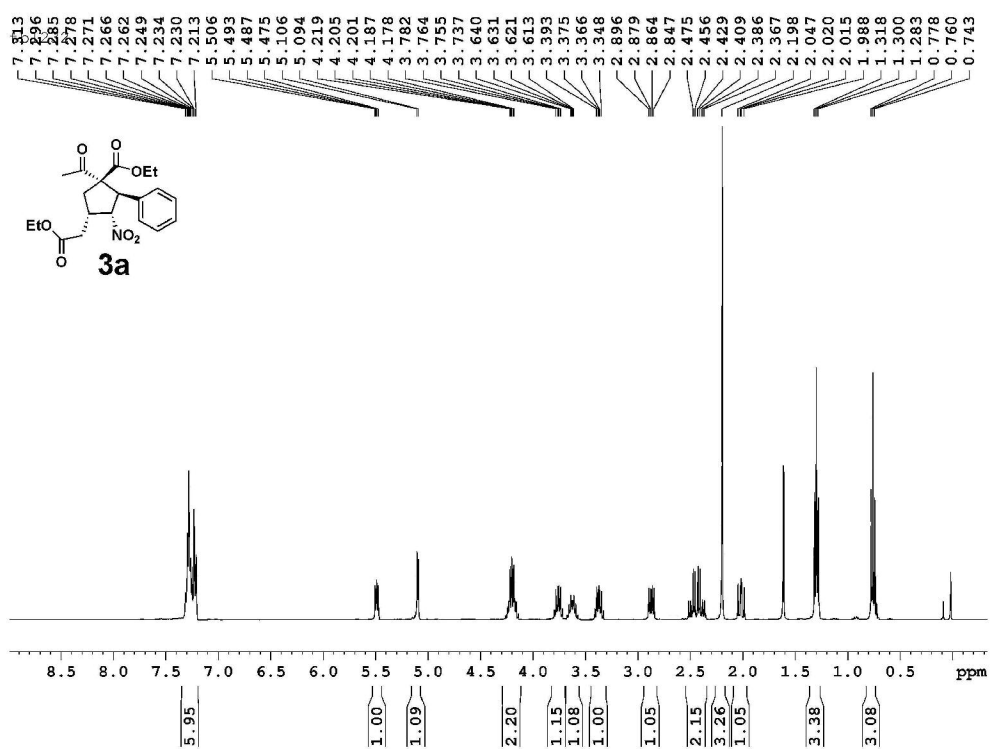
¹³C-NMR (100 MHz, CDCl₃) δ 200.66, 170.89, 170.50, 135.91, 135.07, 134.36, 128.75, 128.73, 128.62, 128.60, 128.13, 126.55, 123.69, 92.81, 69.04, 67.85, 61.03, 51.55, 38.04, 37.63, 34.59, 27.64, 14.15.

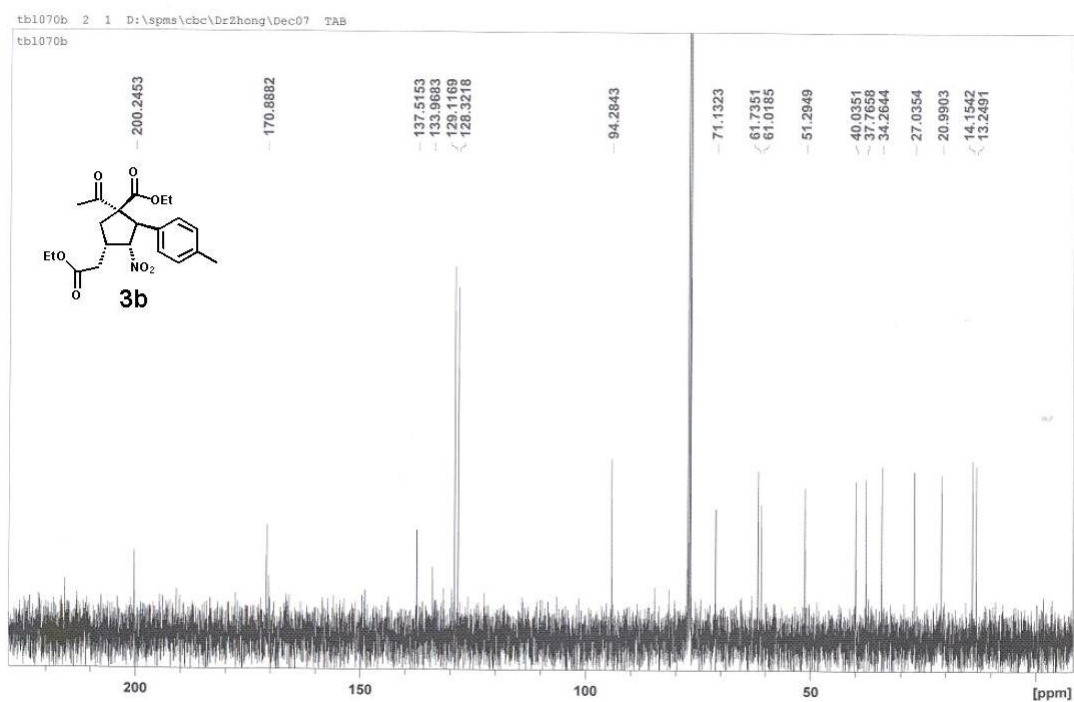
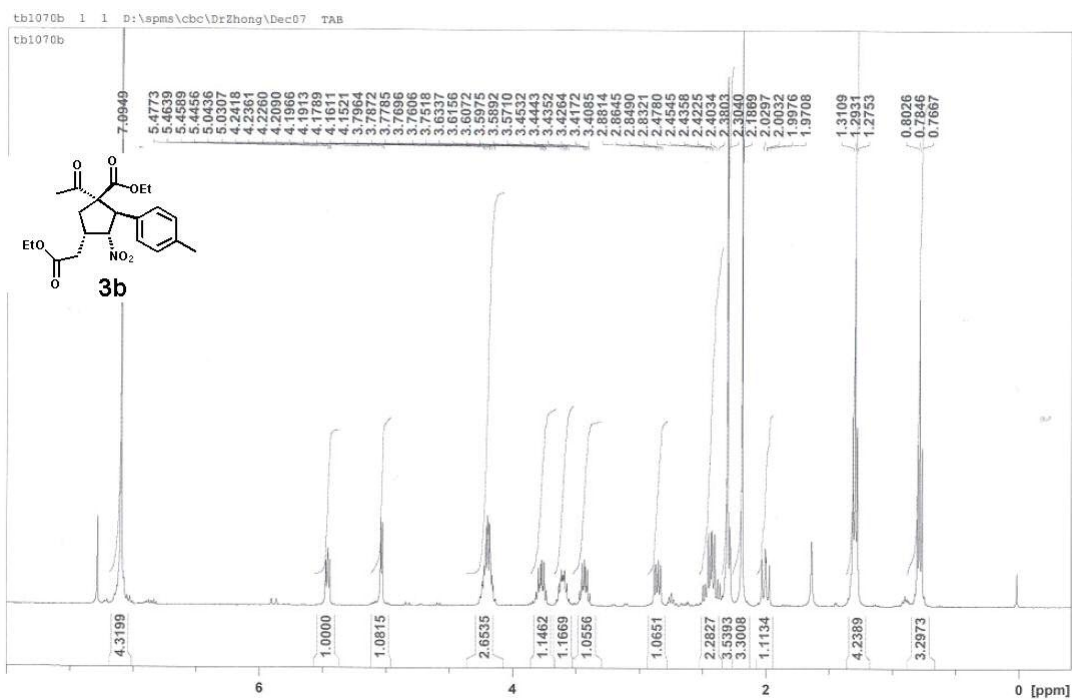
HPLC: Chiralpak AD-H (hexane / *i*-PrOH = 90 / 10, flow rate 1 mL / min, λ = 254 nm), *t_R* (major) = 20.7 min, *t_R* (minor) = 22.8 min; 95% ee.

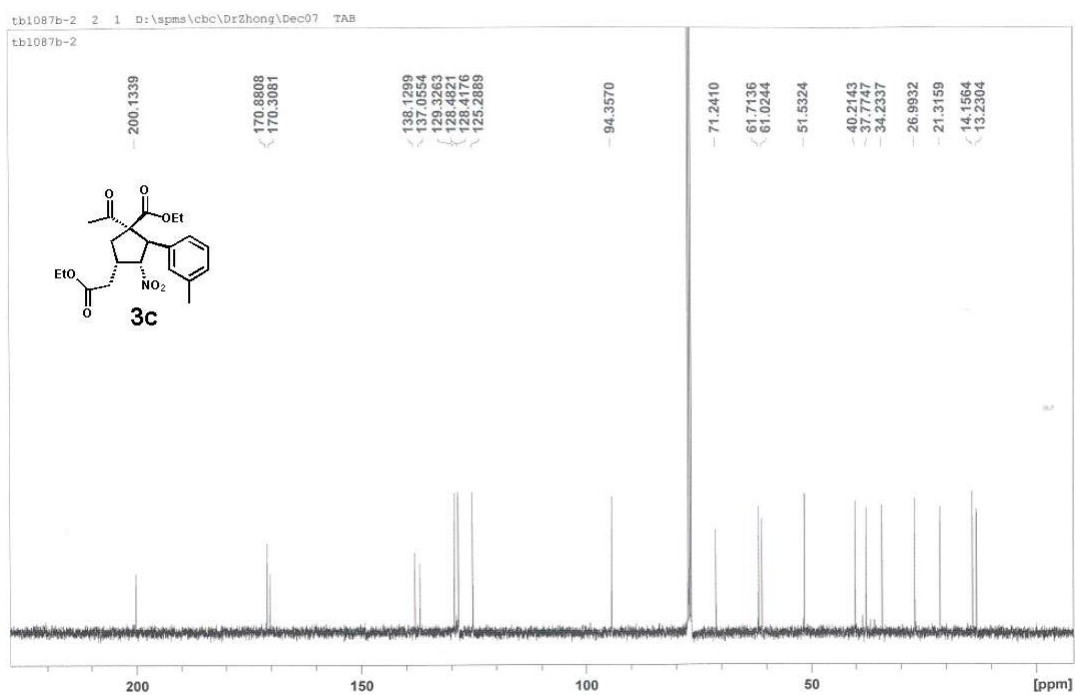
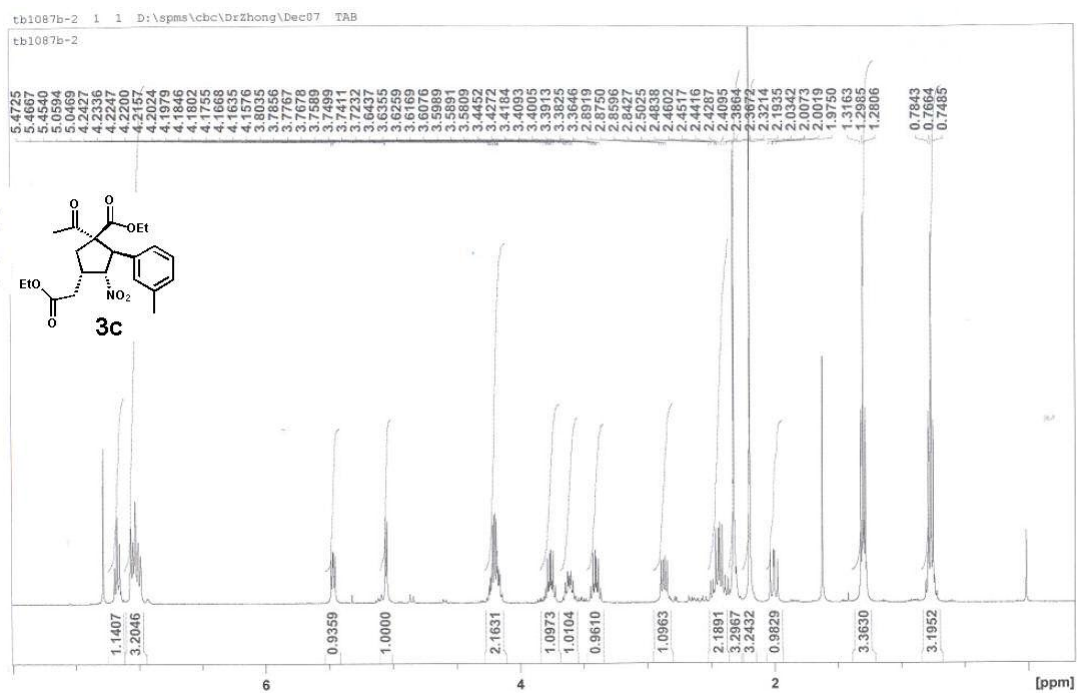
$[\alpha]_{\text{D}}^{22} = -5.3$ ($c = 1.0$, CH_2Cl_2).

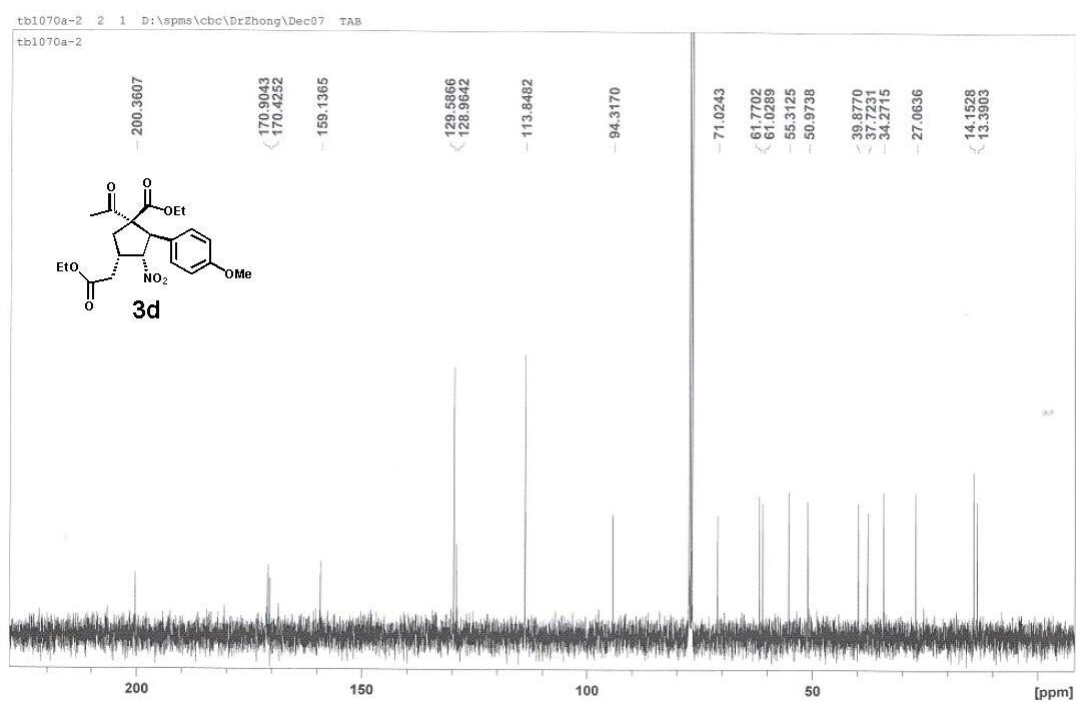
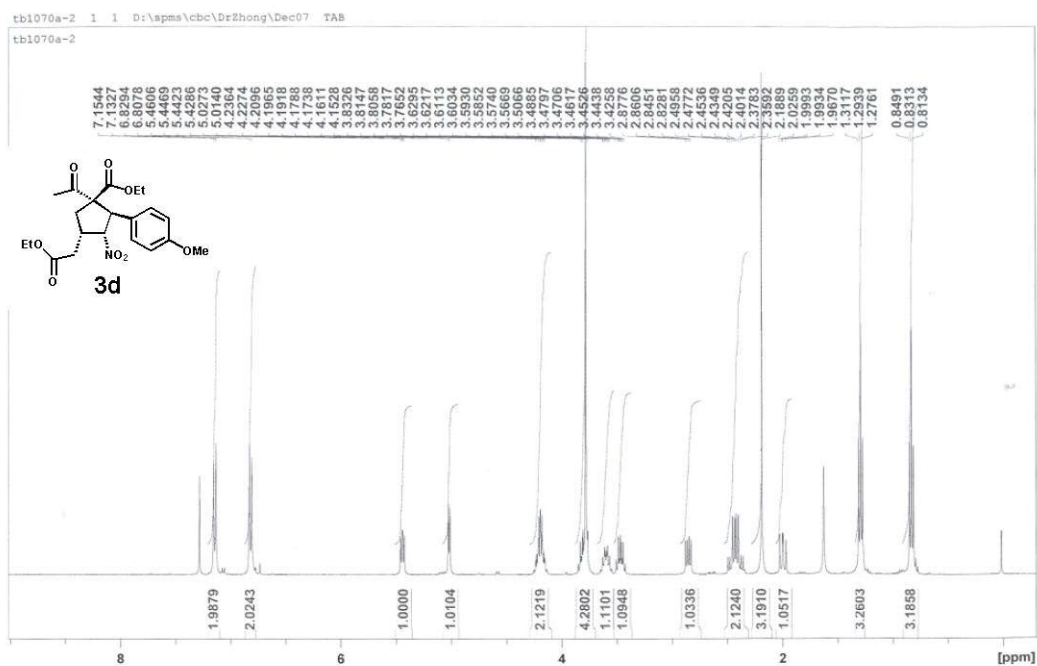
HRMS (ESI) calcd for $\text{C}_{27}\text{H}_{29}\text{NO}_7 + \text{H}$, m/z 480.2022, found 480.2017.

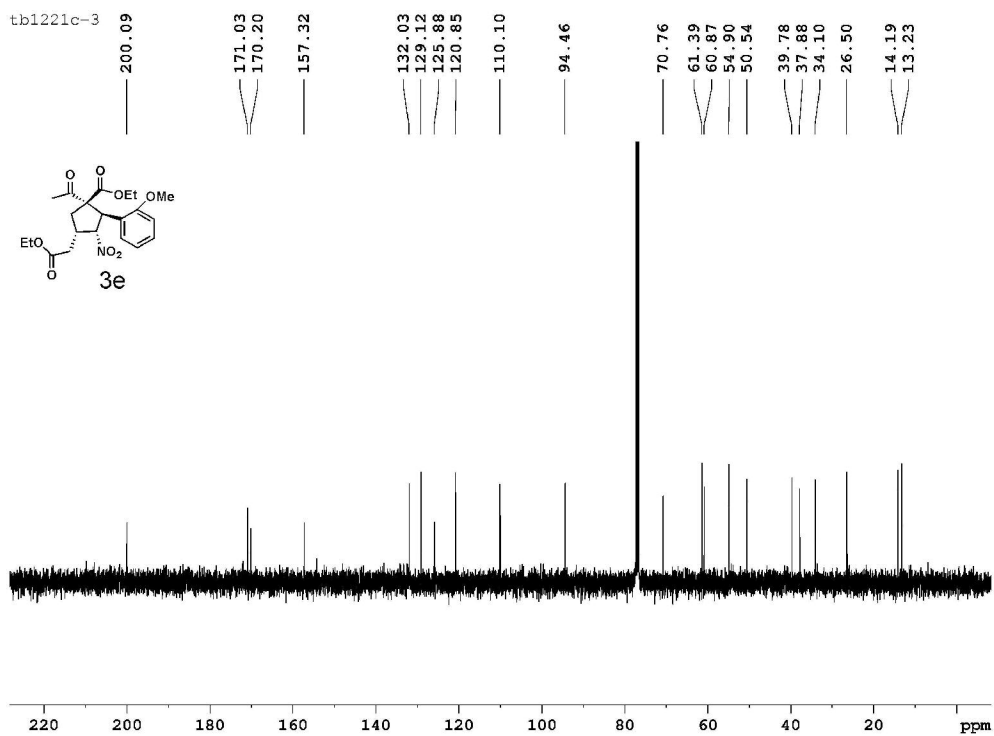
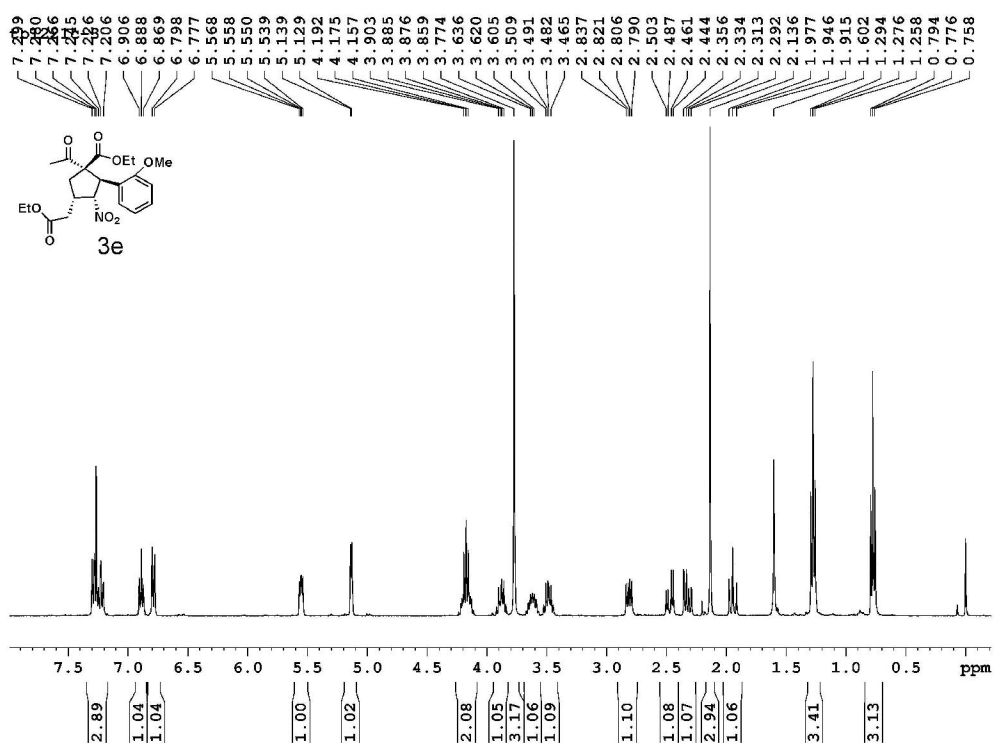
NMR Spectra

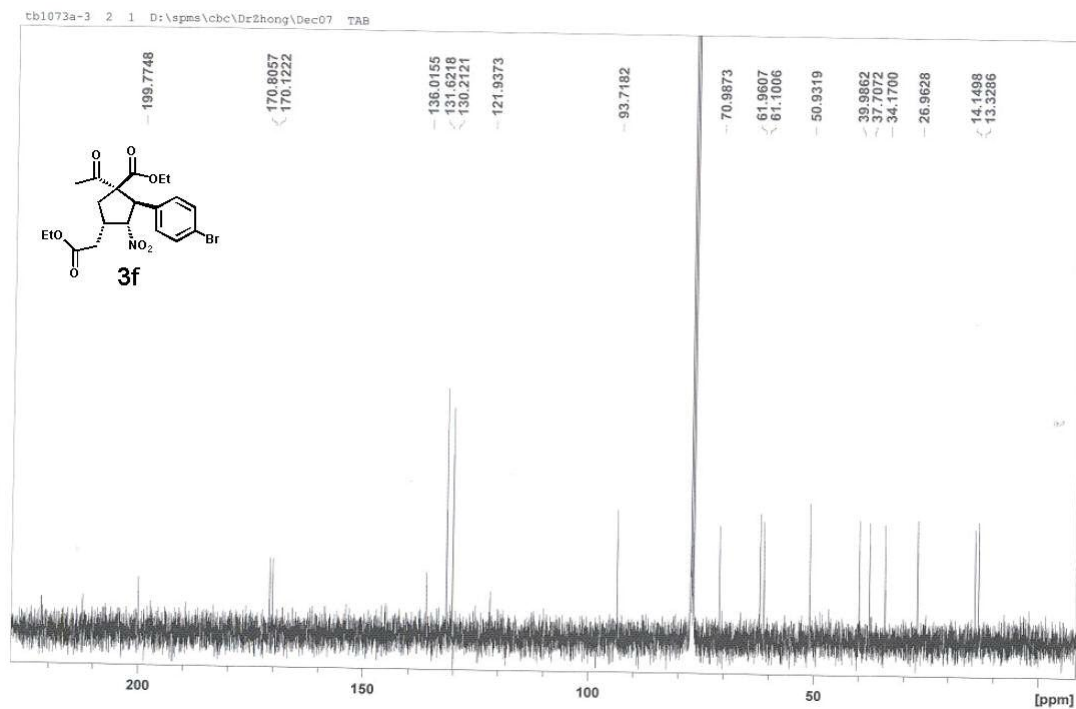
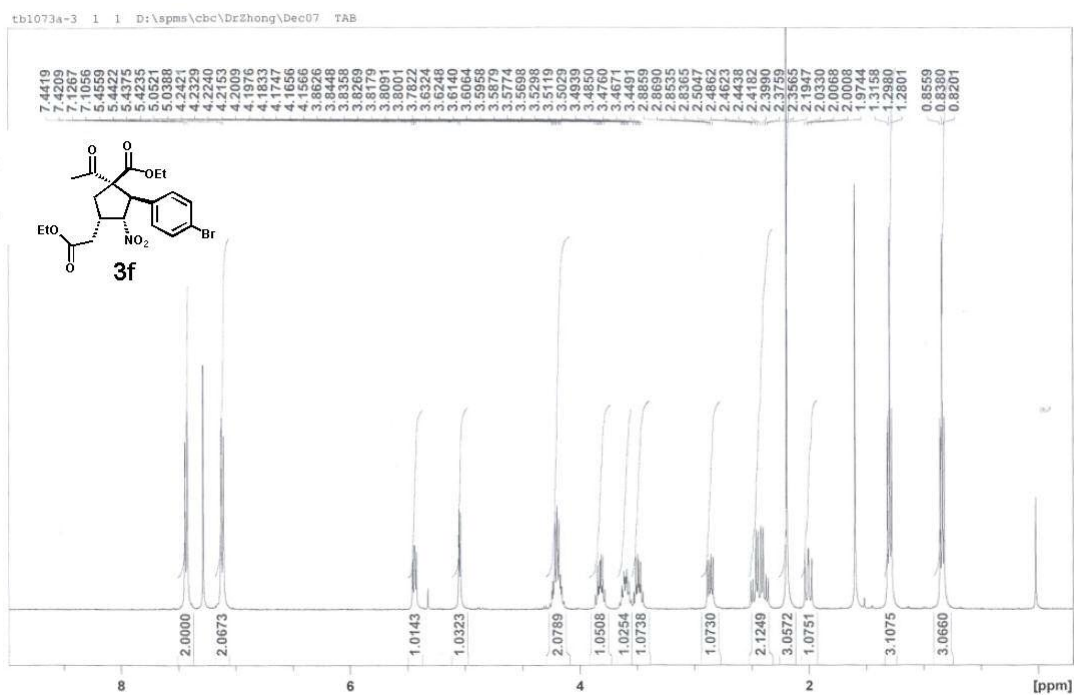


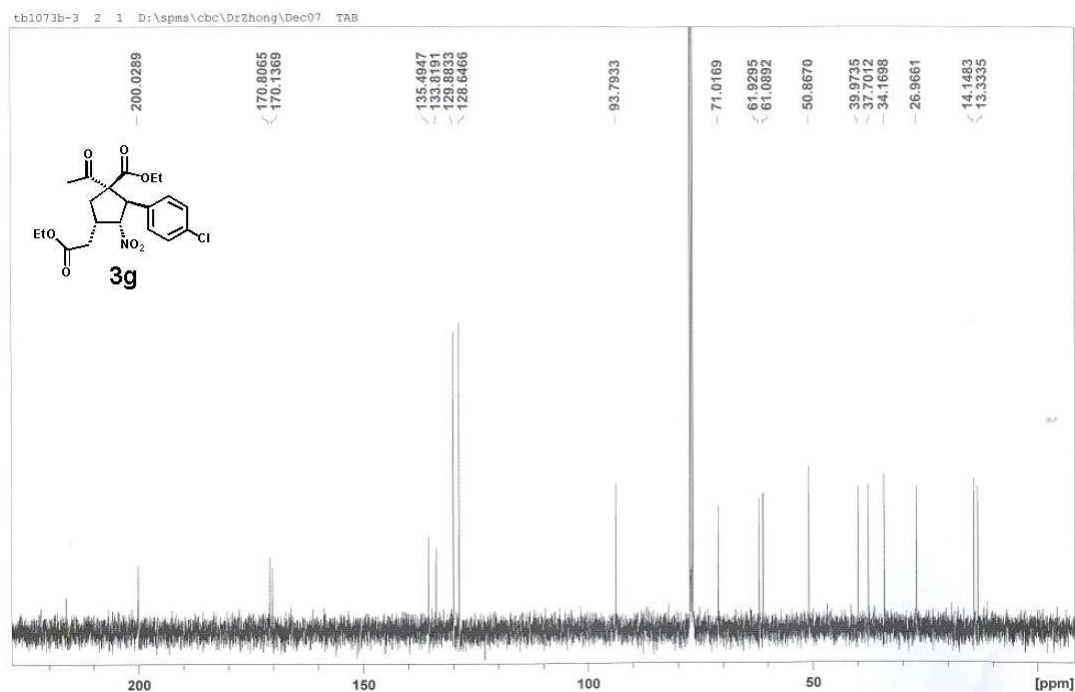
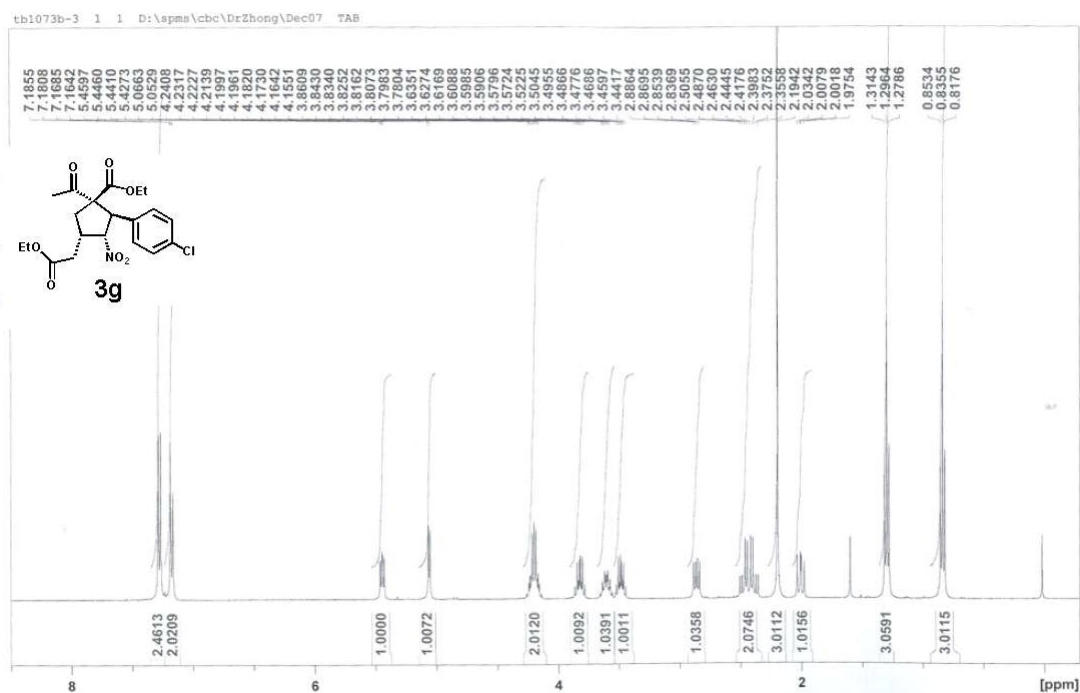




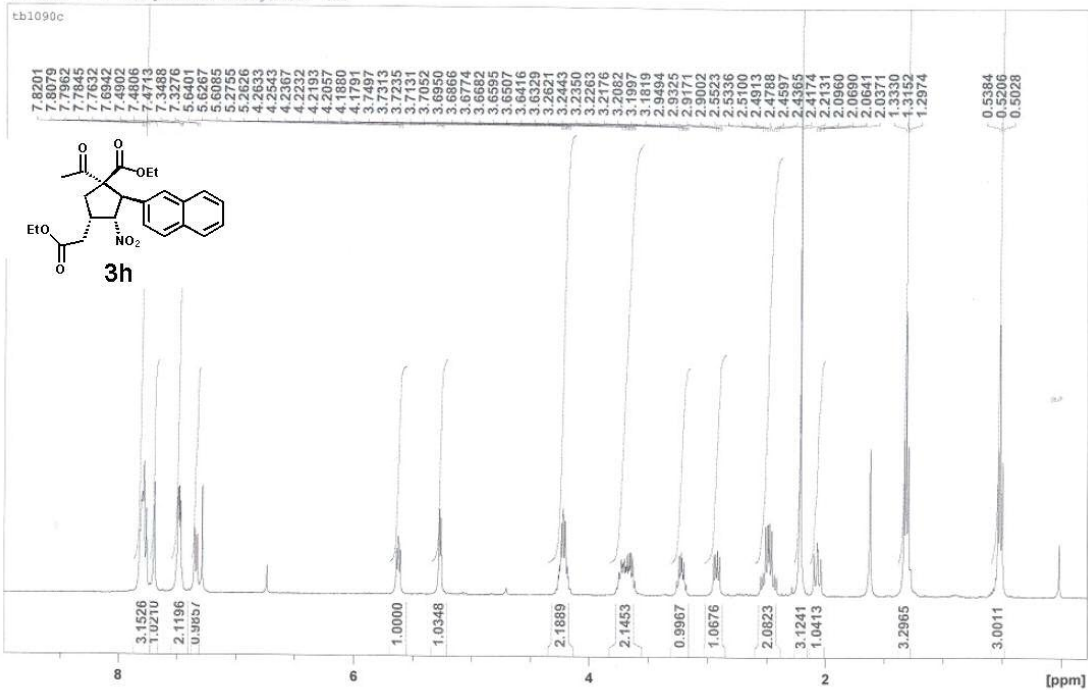




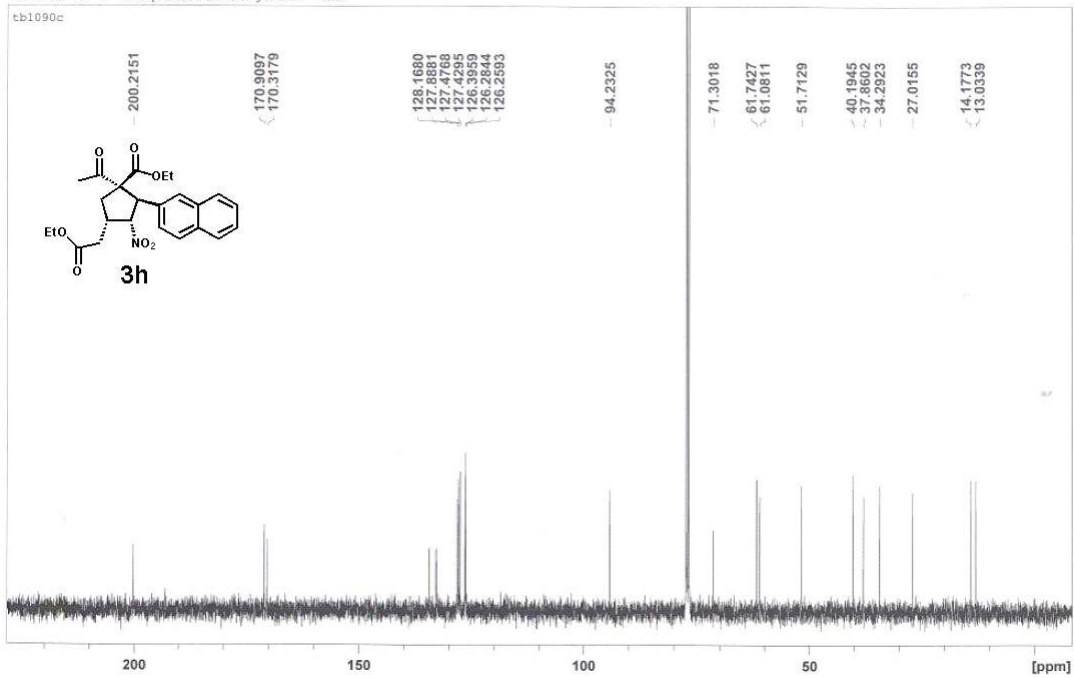


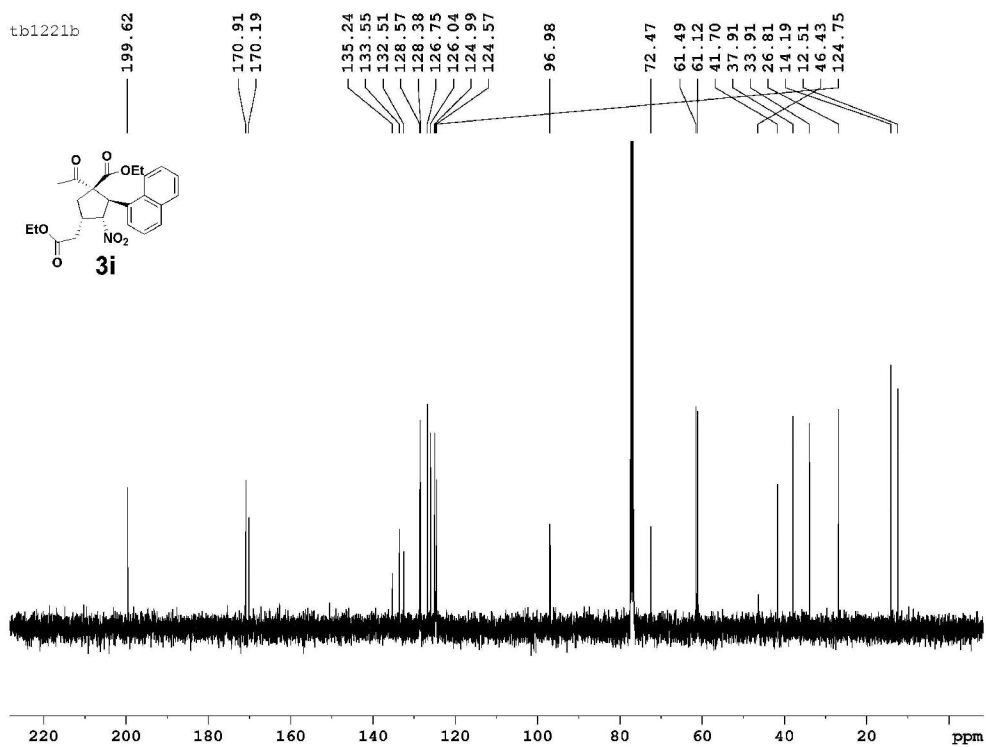
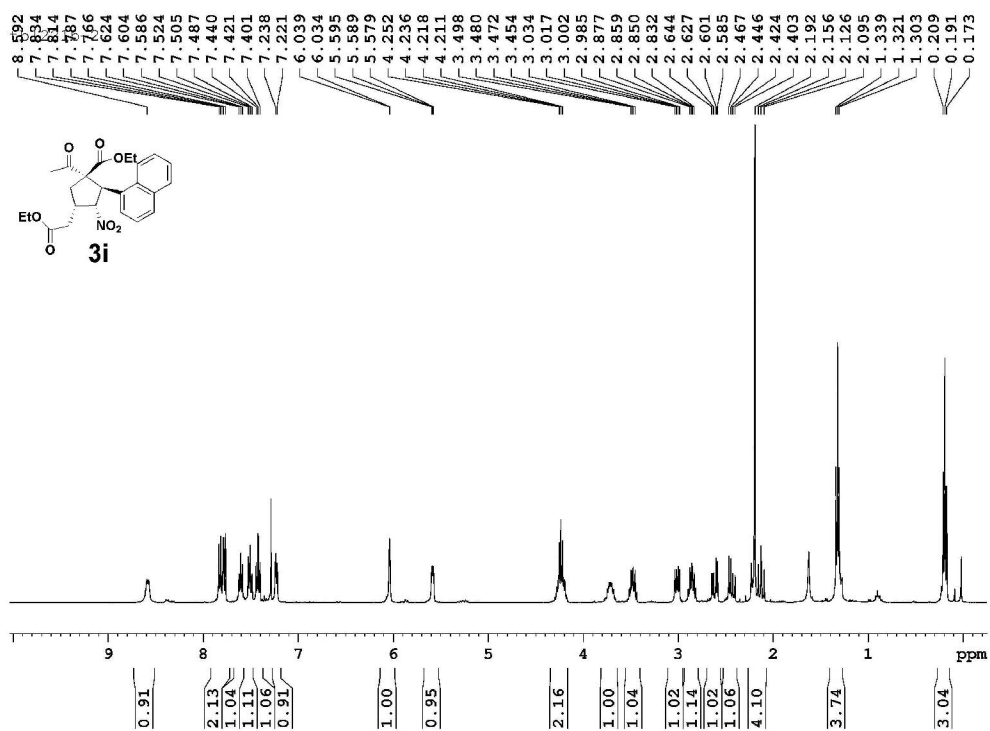


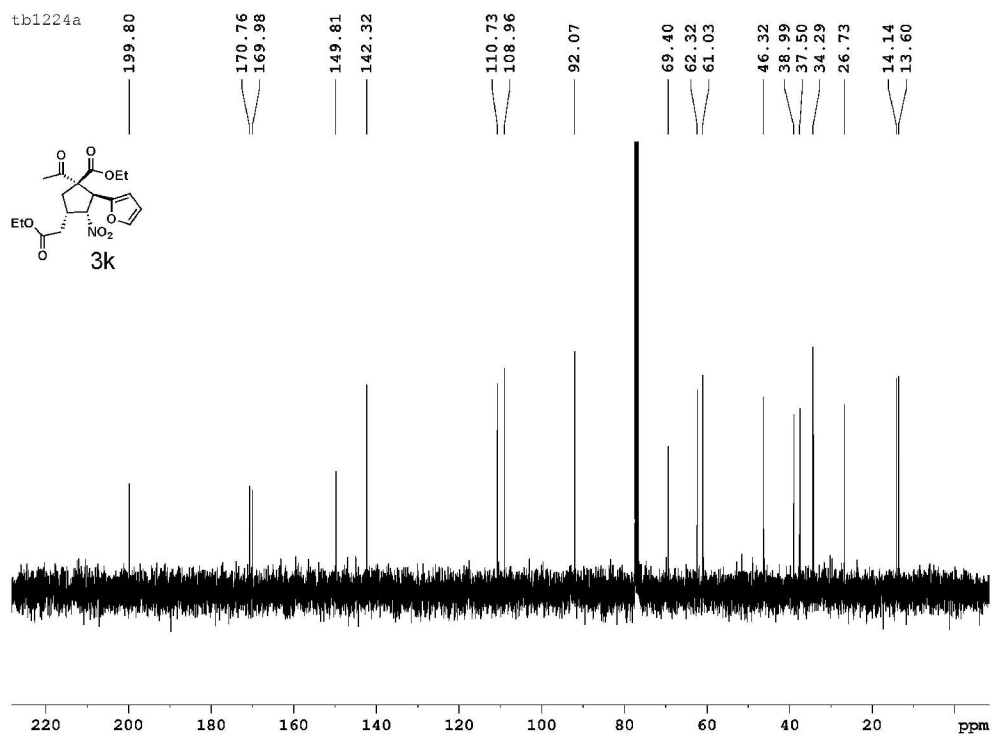
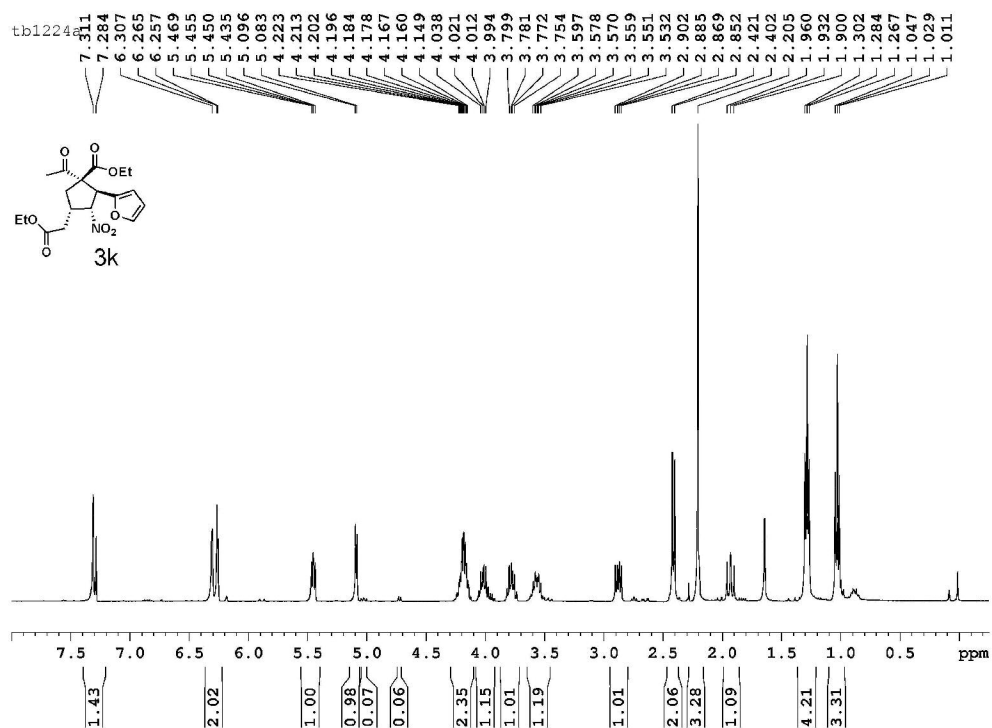
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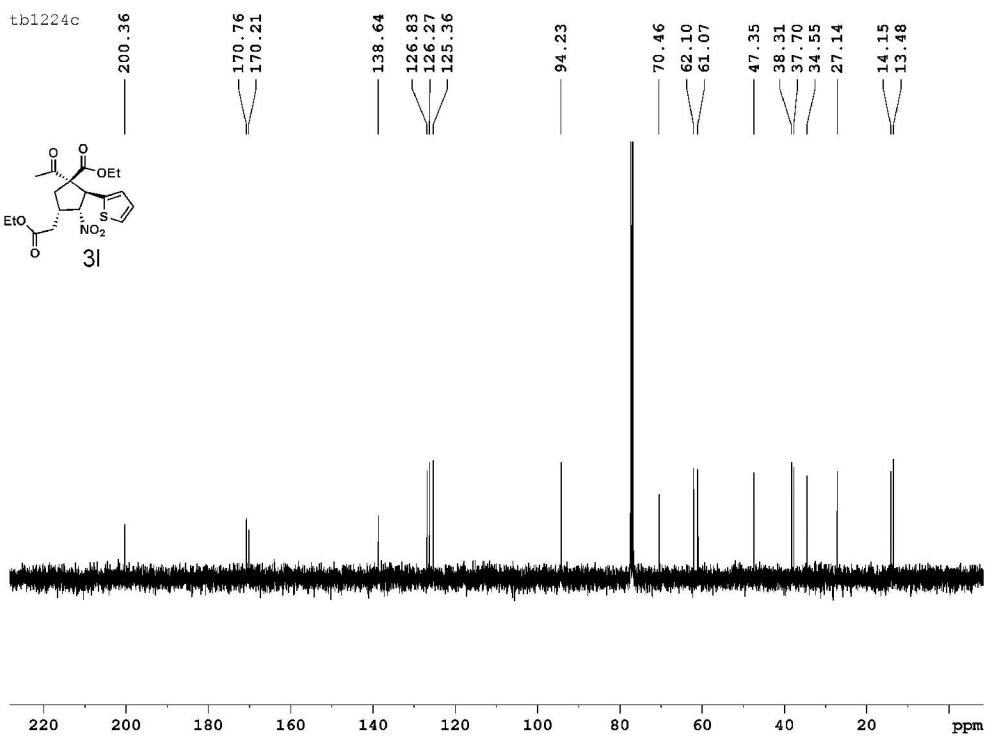
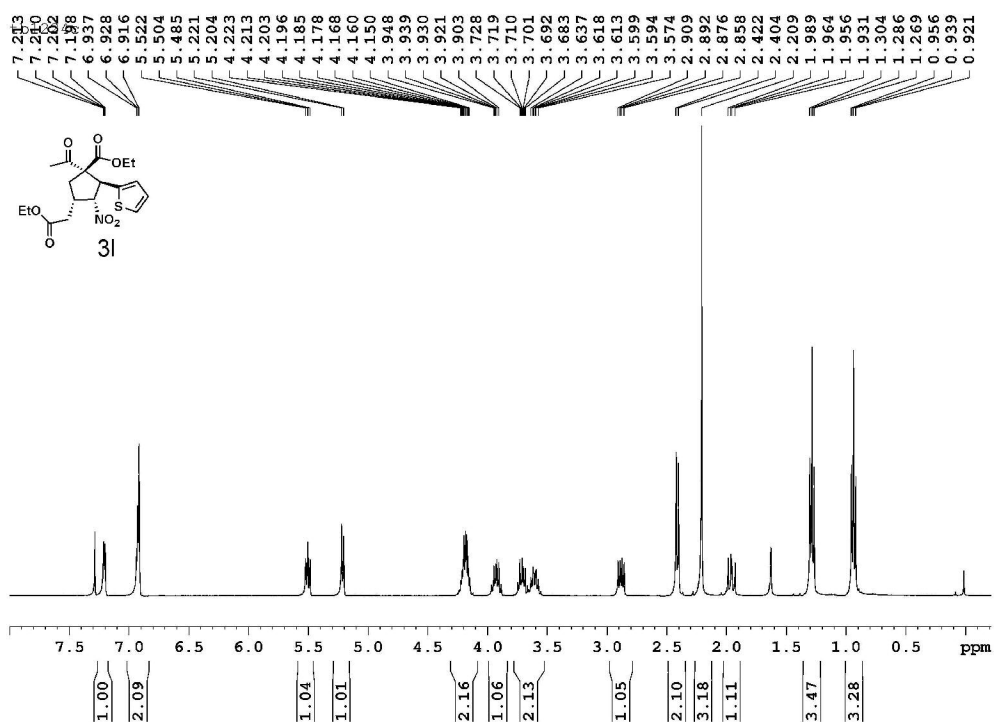


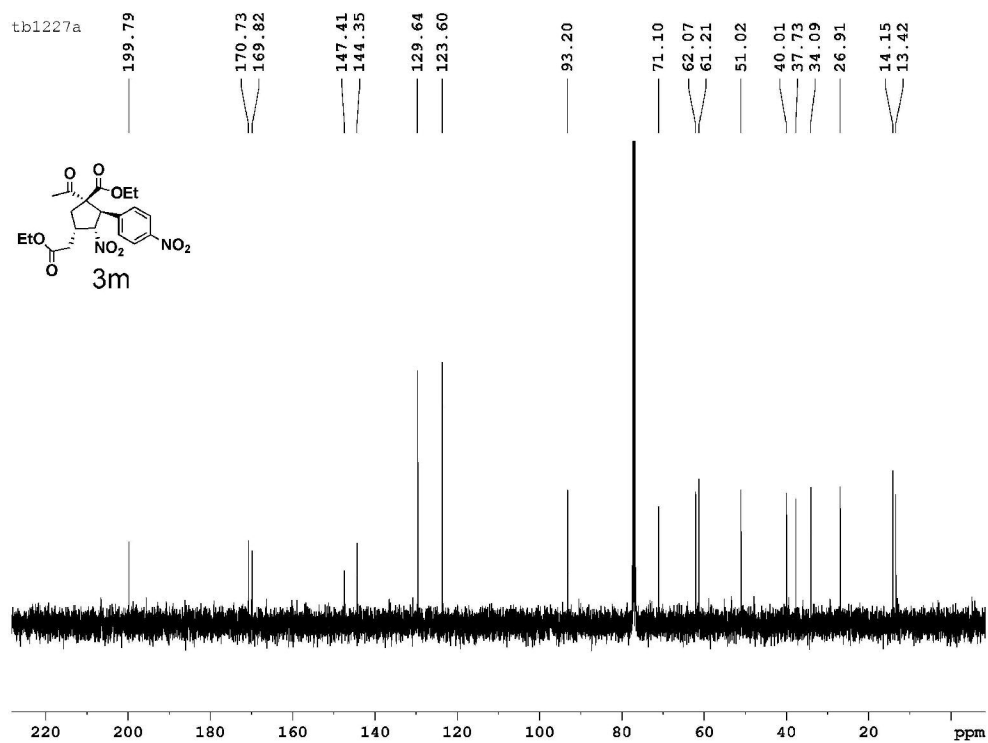
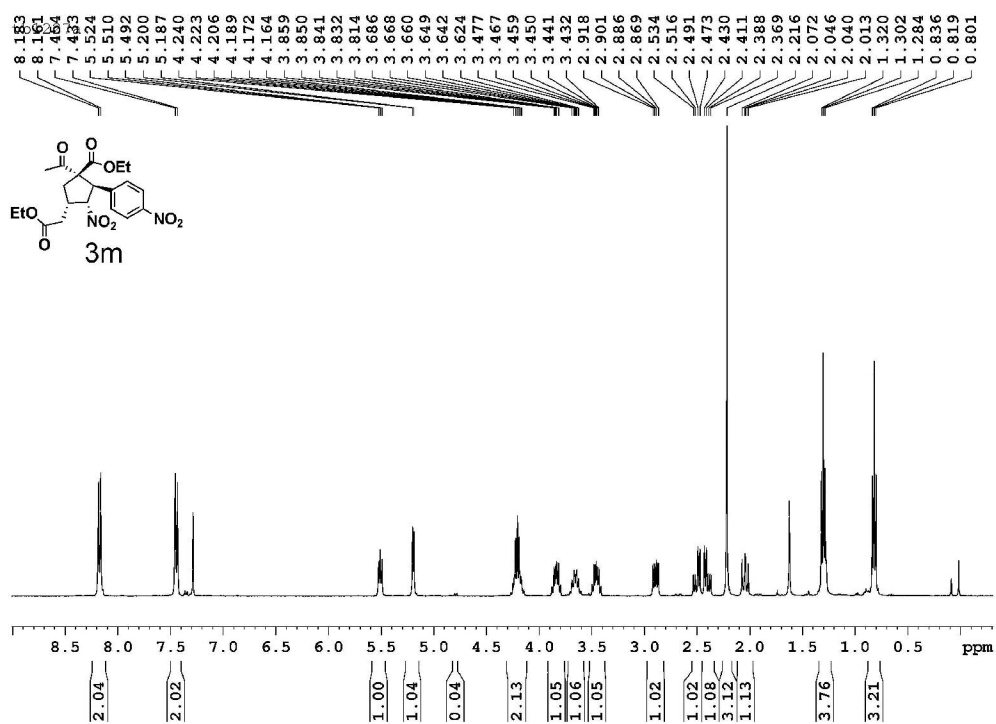
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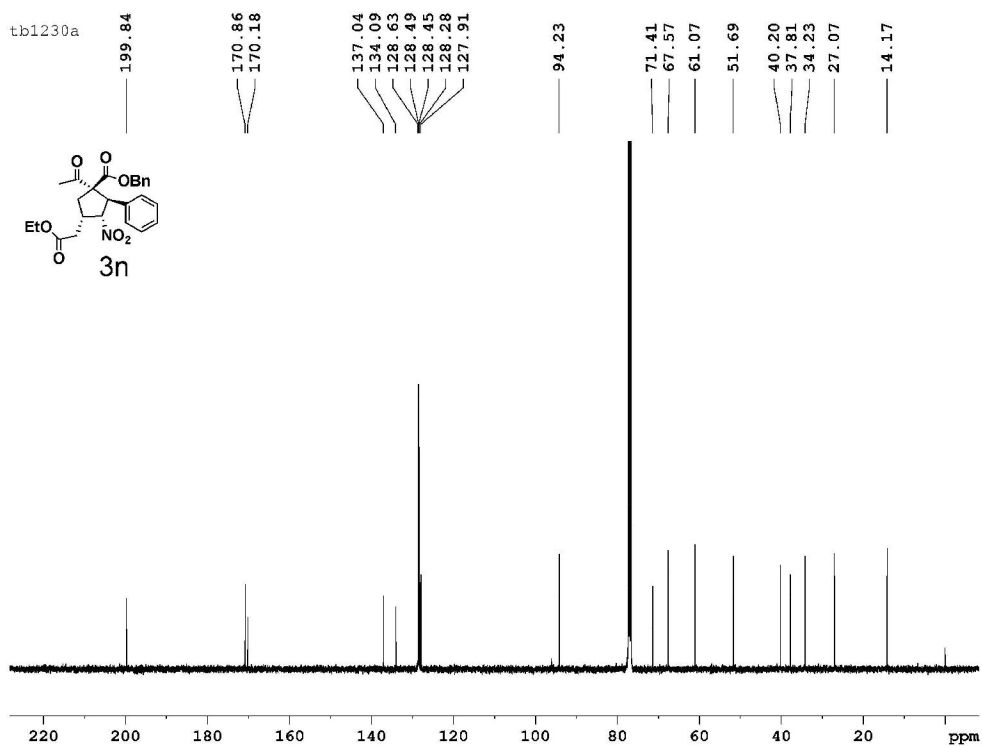
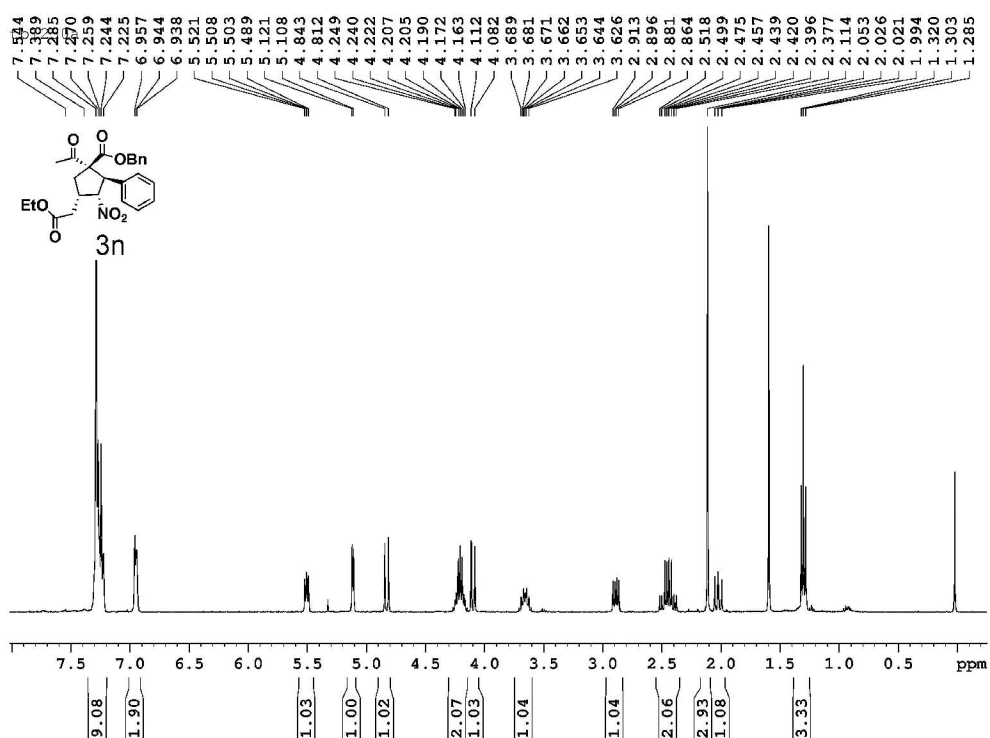


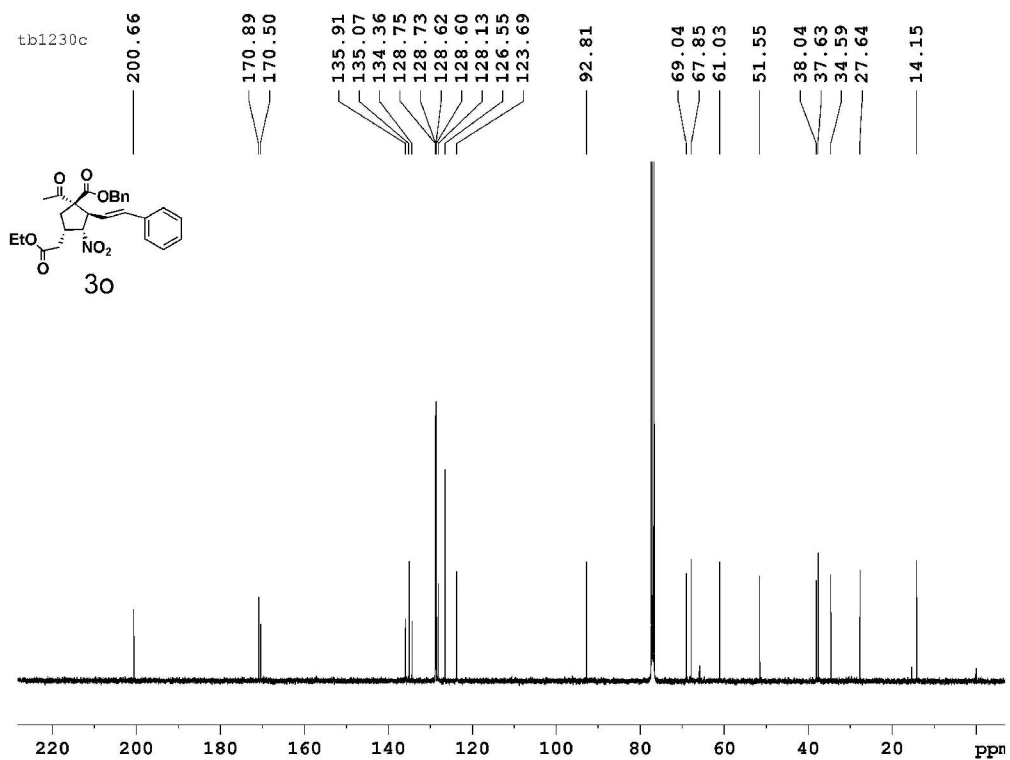
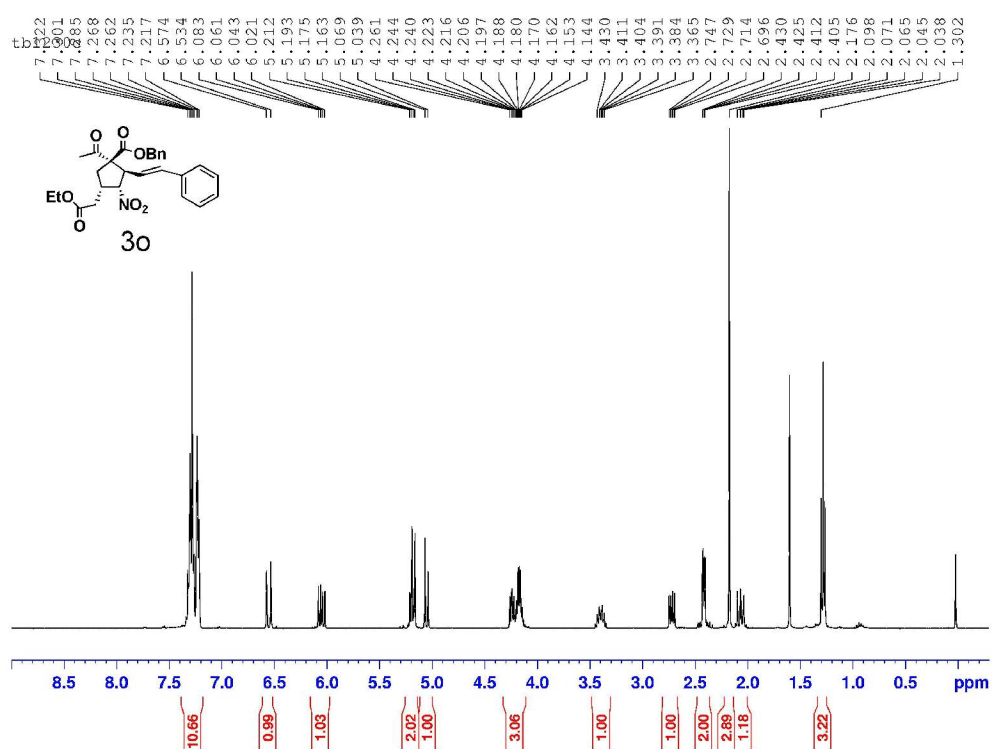




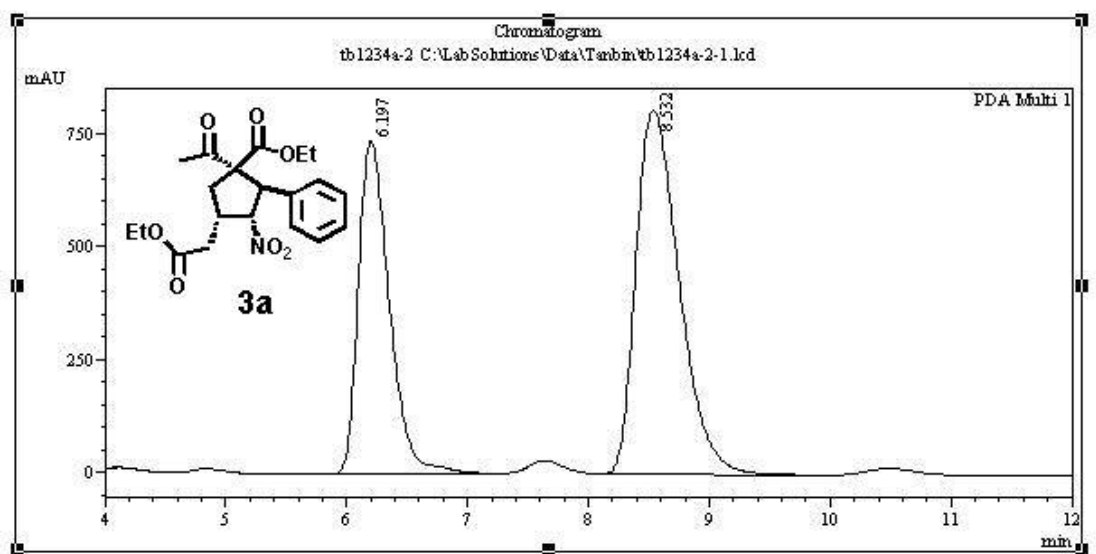








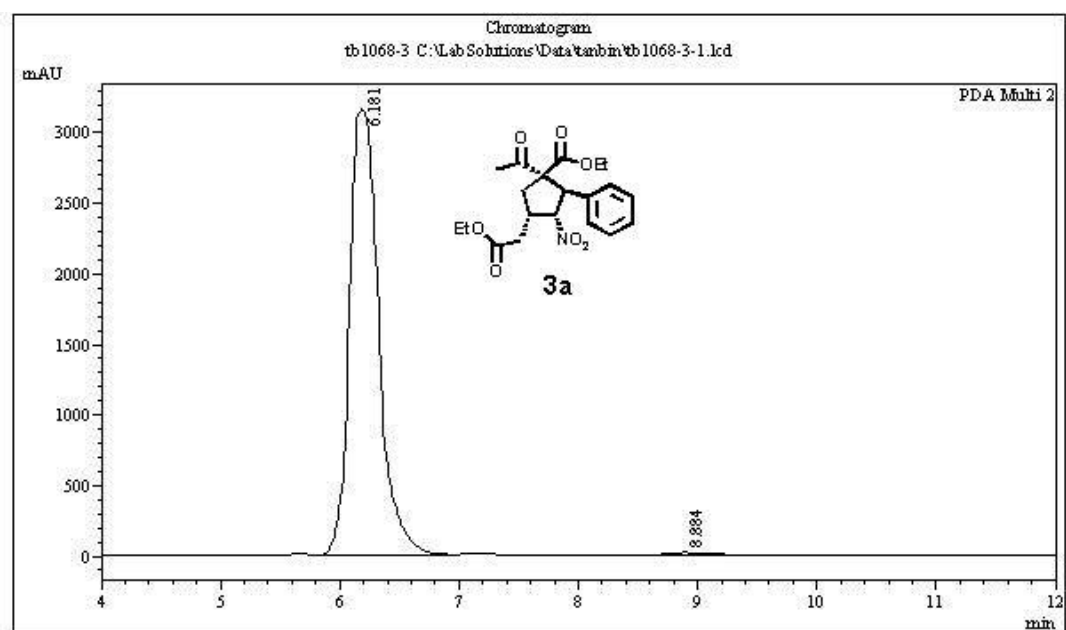
HPLC Spectra



PeakTable

PDA Ch1 220nm 4nm

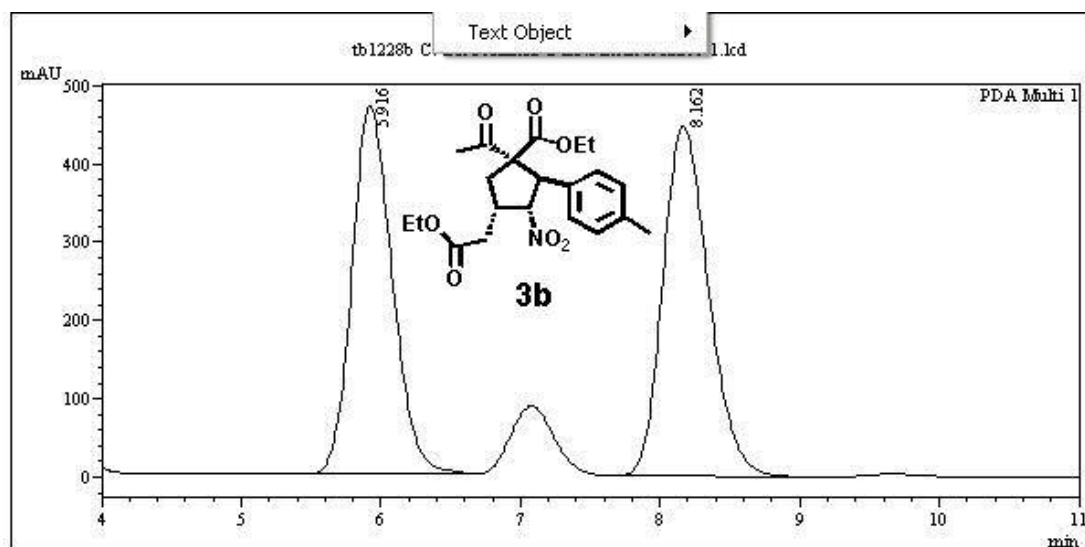
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PeakTable

PDA Ch2 230nm 4nm

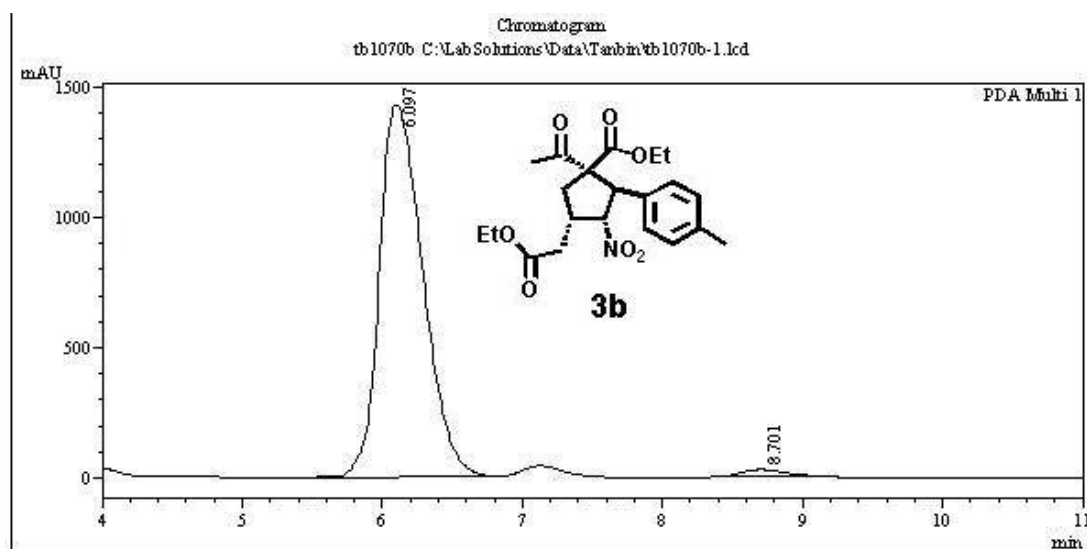
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PeakTable

PDA Ch1 220nm 4nm

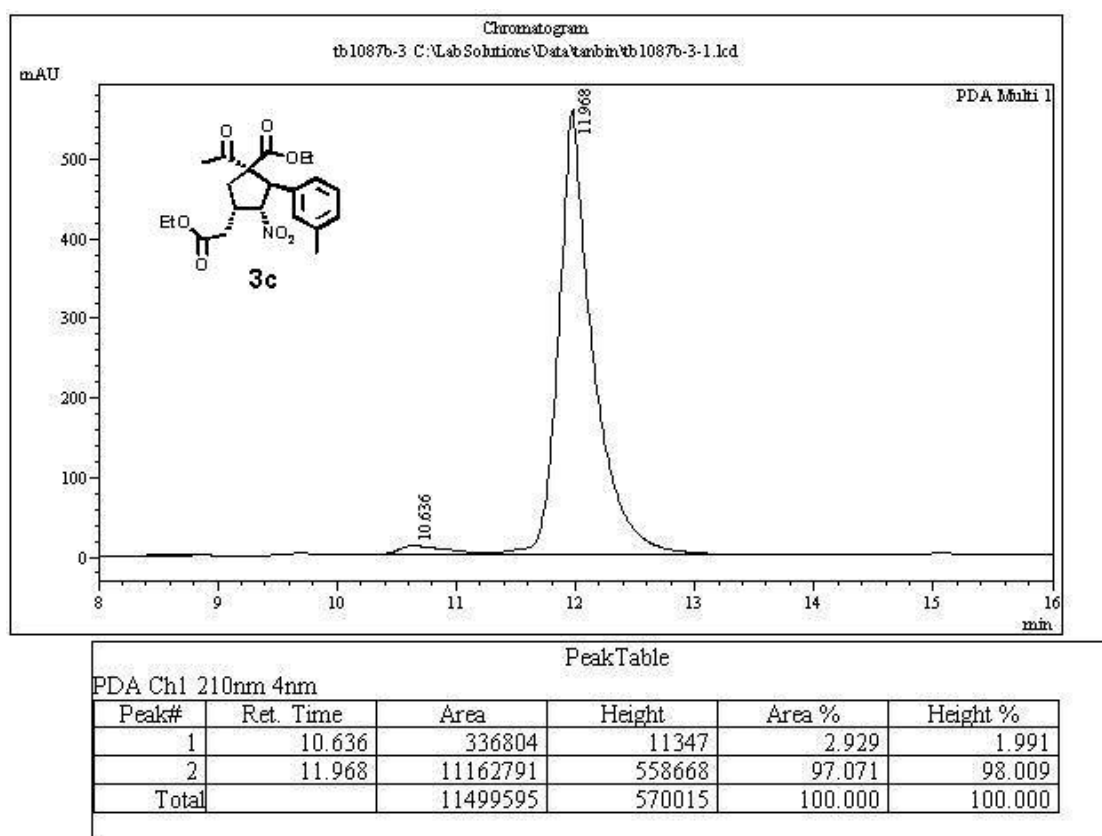
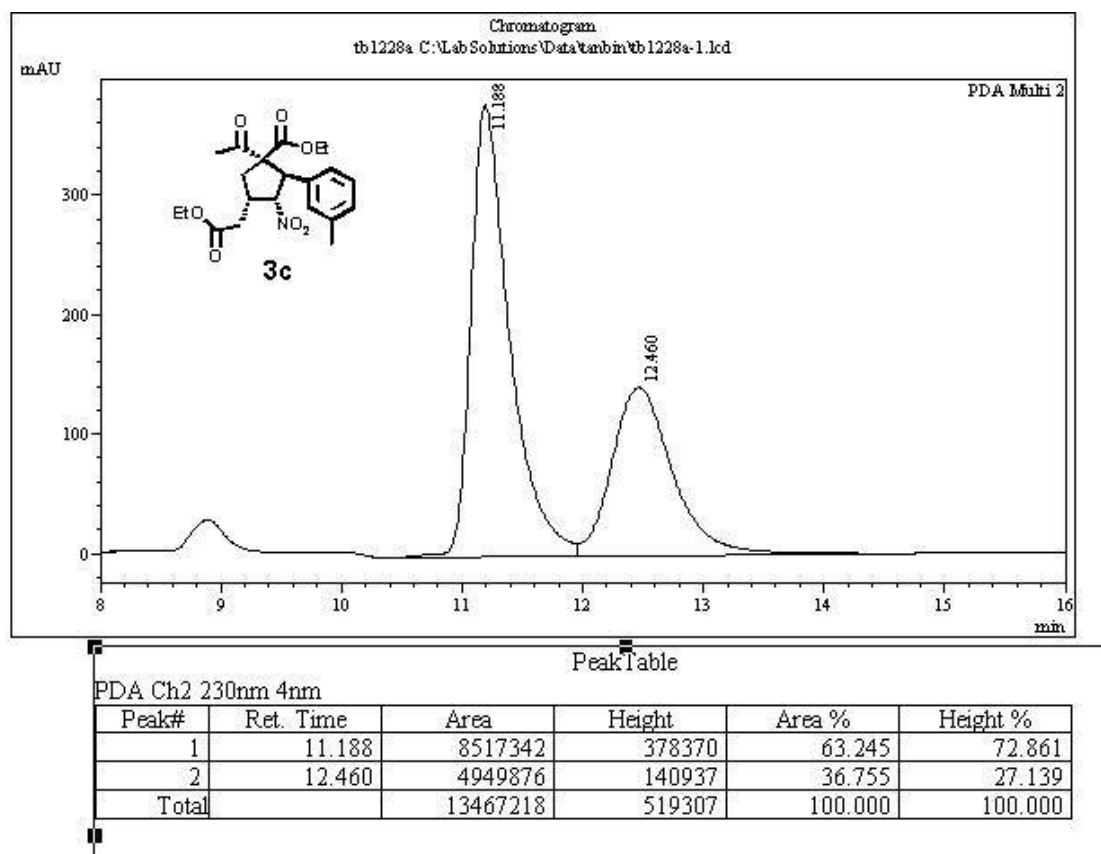
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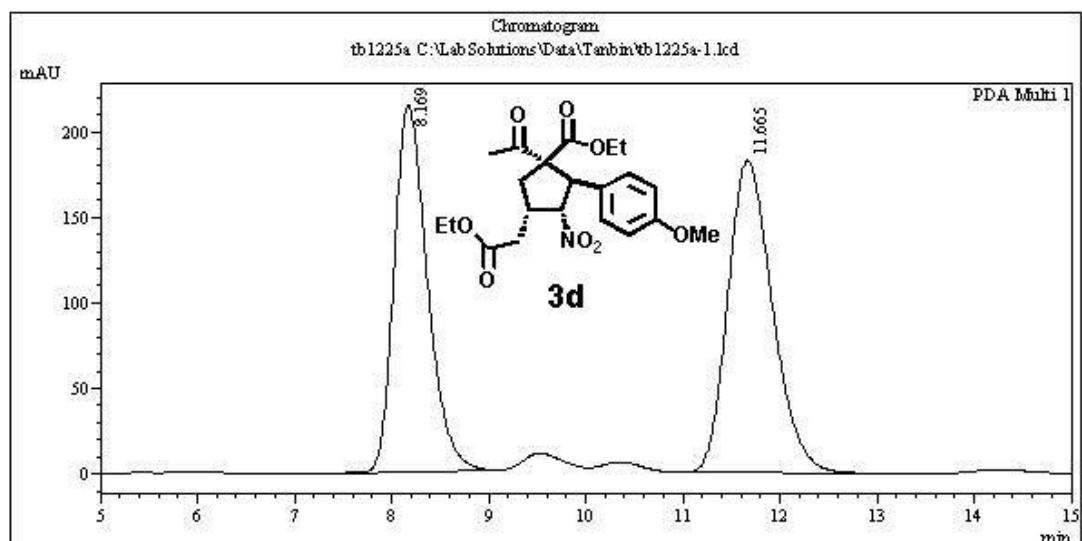


PeakTable

PDA Ch1 220nm 4nm

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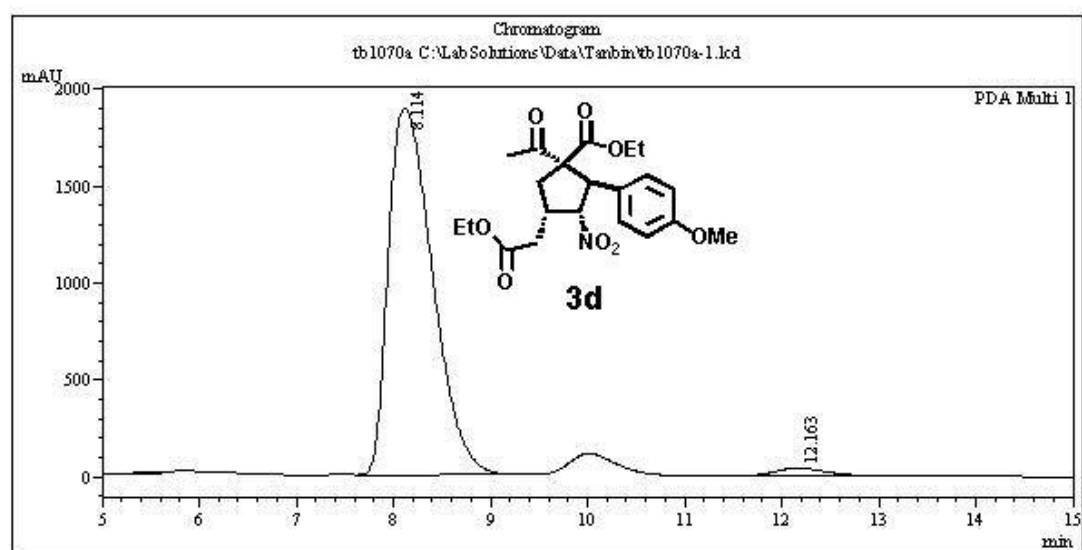




PeakTable

PDA Ch1 220nm 4nm

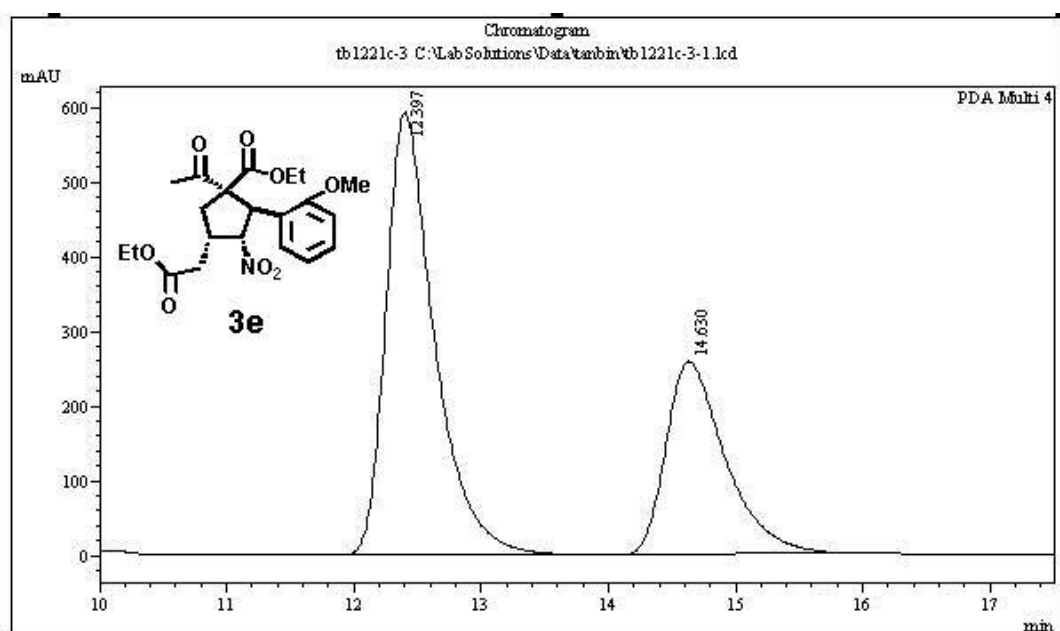
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PeakTable

PDA Ch1 220nm 4nm

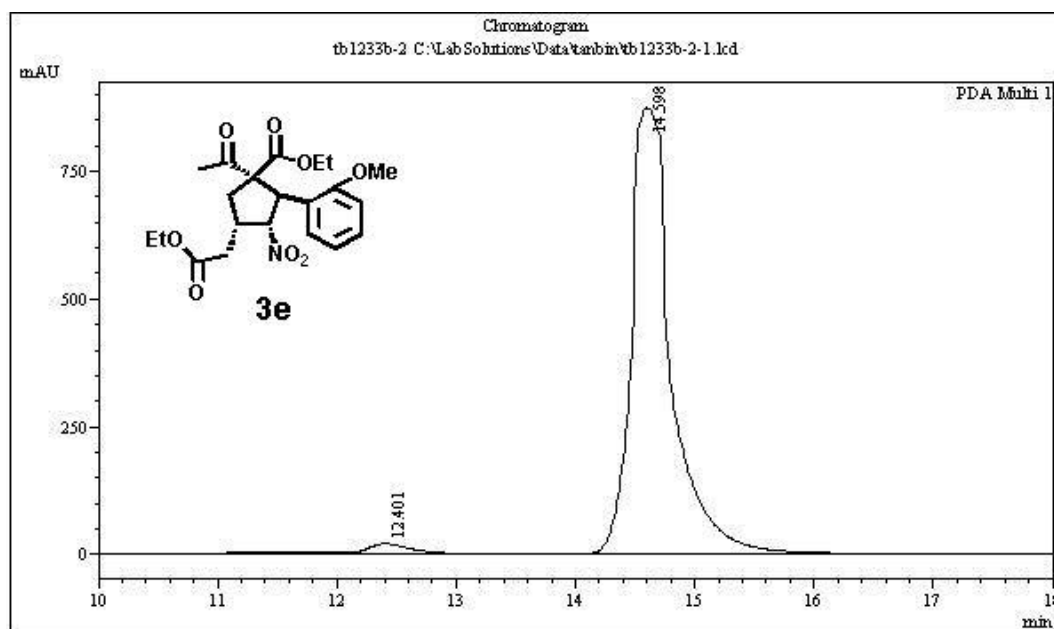
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PeakTable

PDA Ch4 220nm 4nm

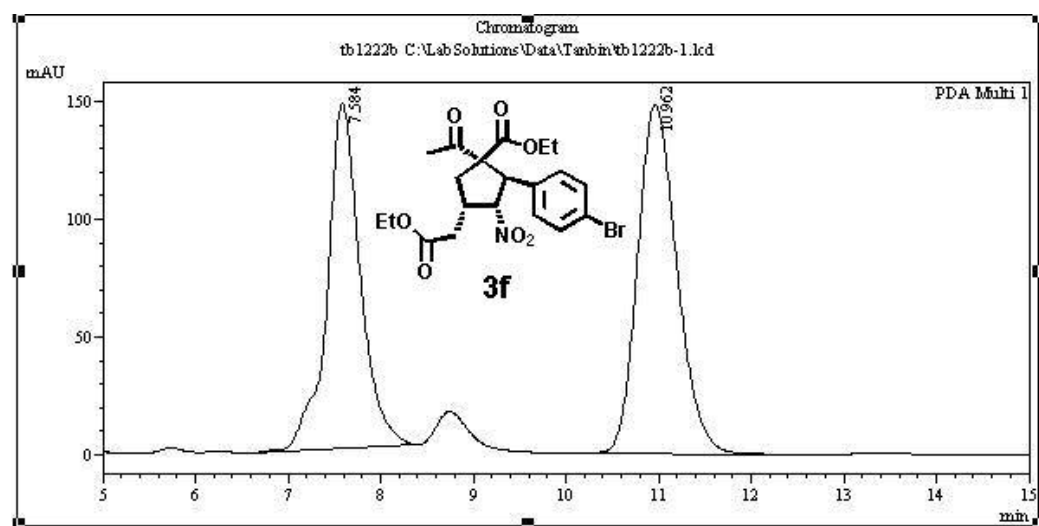
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PeakTable

PDA Ch1 210nm 4nm

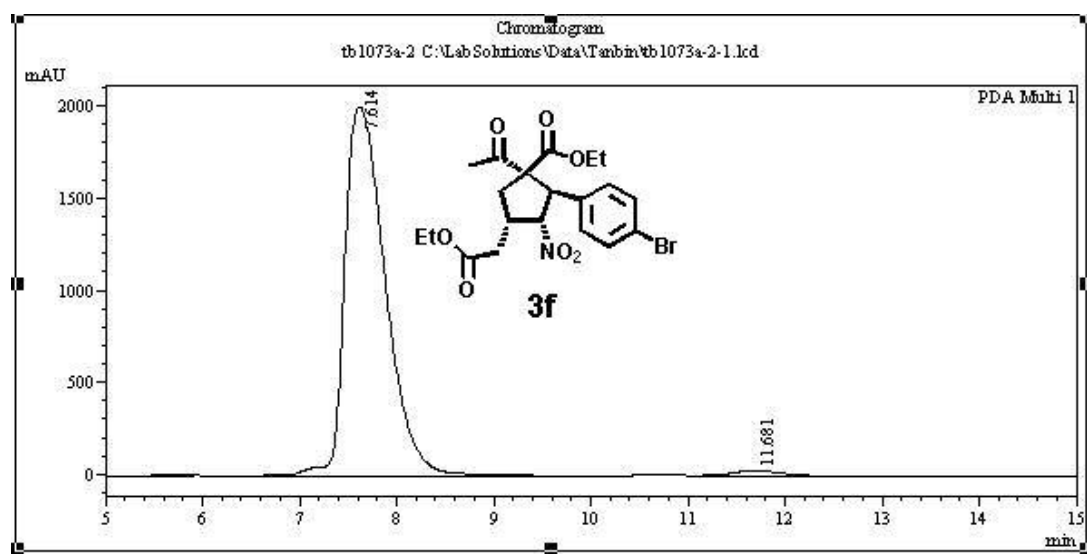
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PeakTable

PDA Ch1 220nm 4nm

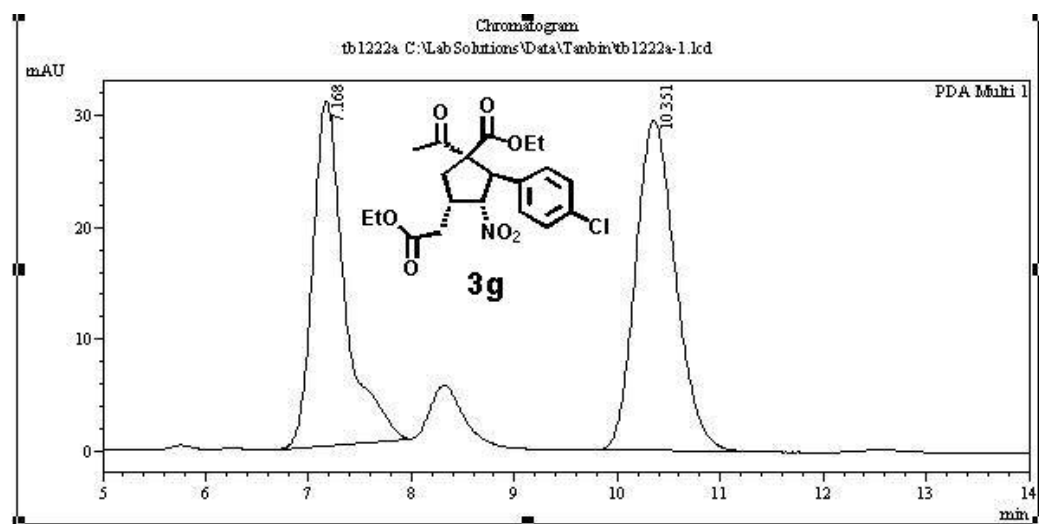
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PeakTable

PDA Ch1 220nm 4nm

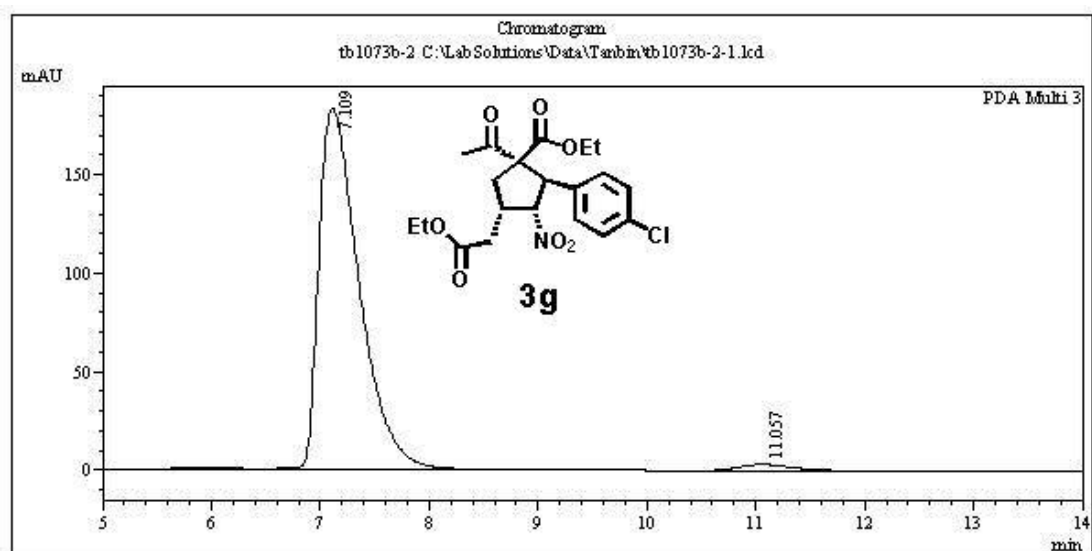
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PeakTable

PDA Ch1 220nm 4nm

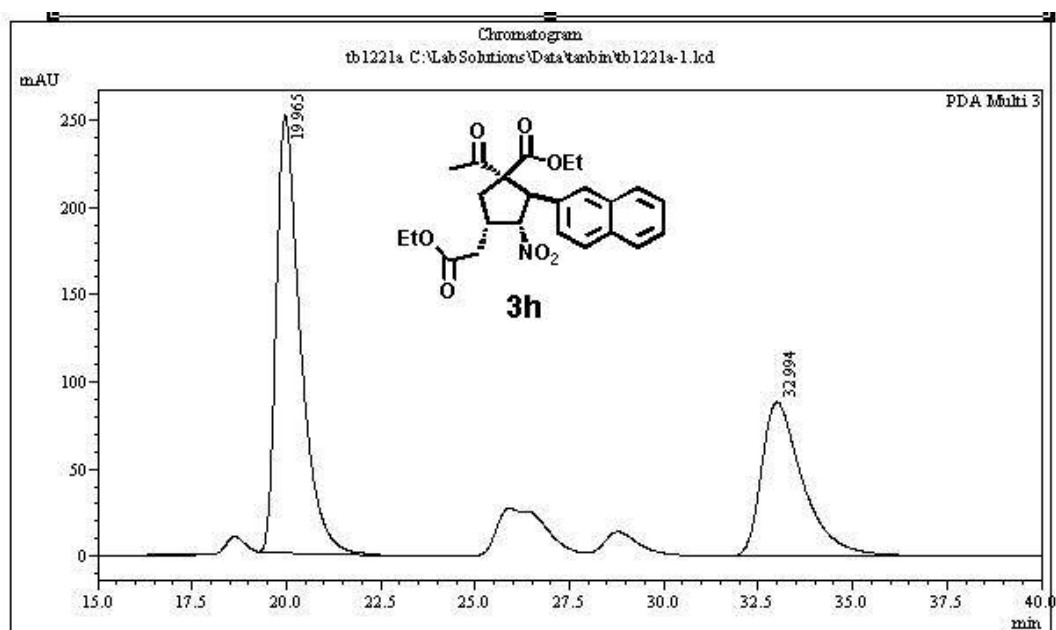
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1	7.168	662374	30710	45.380	51.037
2	10.351	797250	29463	54.620	48.963
Total		1459625	60173	100.000	100.000



PeakTable

PDA Ch3 254nm 4nm

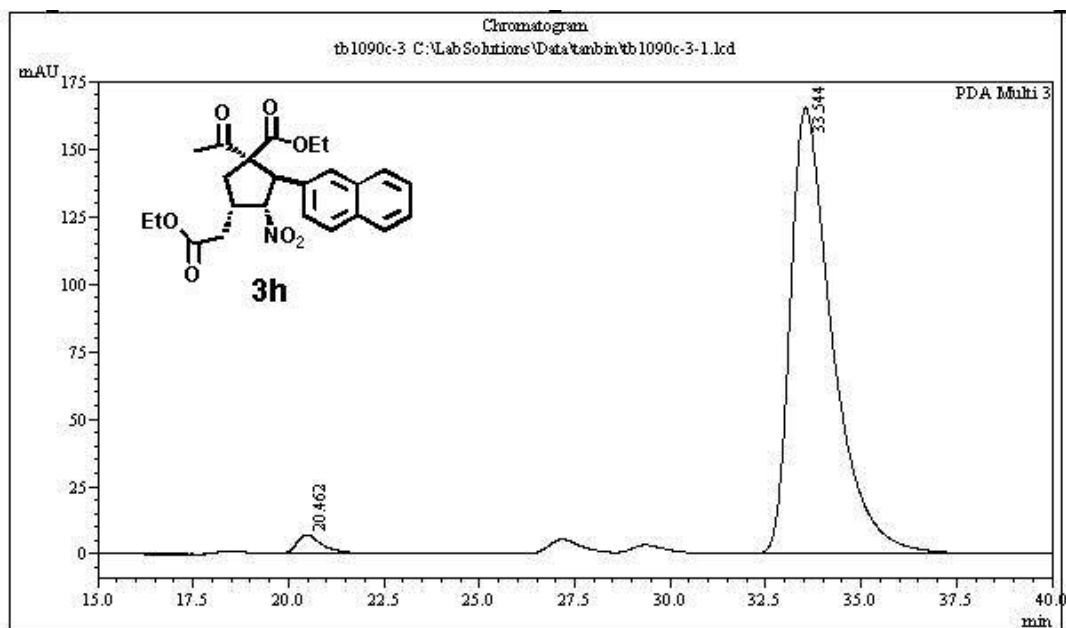
Peak#	Ret. Time	Area	Height	Area %	Height %
1	7.109	4725021	183370	97.860	98.327
2	11.057	103310	3119	2.140	1.673
Total		4828331	186489	100.000	100.000



PeakTable

PDA Ch3 254nm 4nm

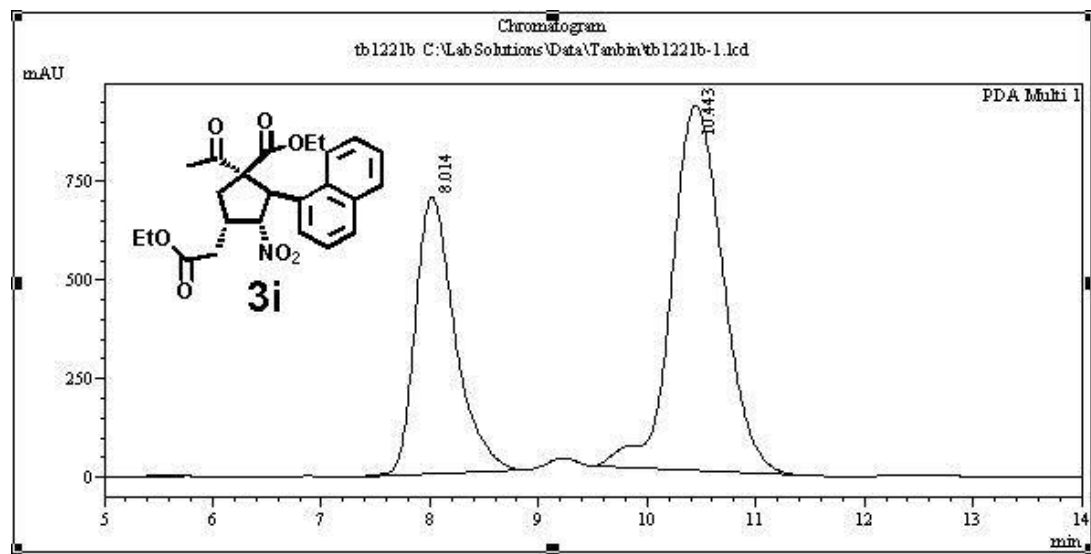
Peak#	Ret. Time	Area	Height	Area %	Height %
1	19.965	11170366	250515	62.705	73.954
2	32.994	6643774	88230	37.295	26.046
Total		17814140	338745	100.000	100.000



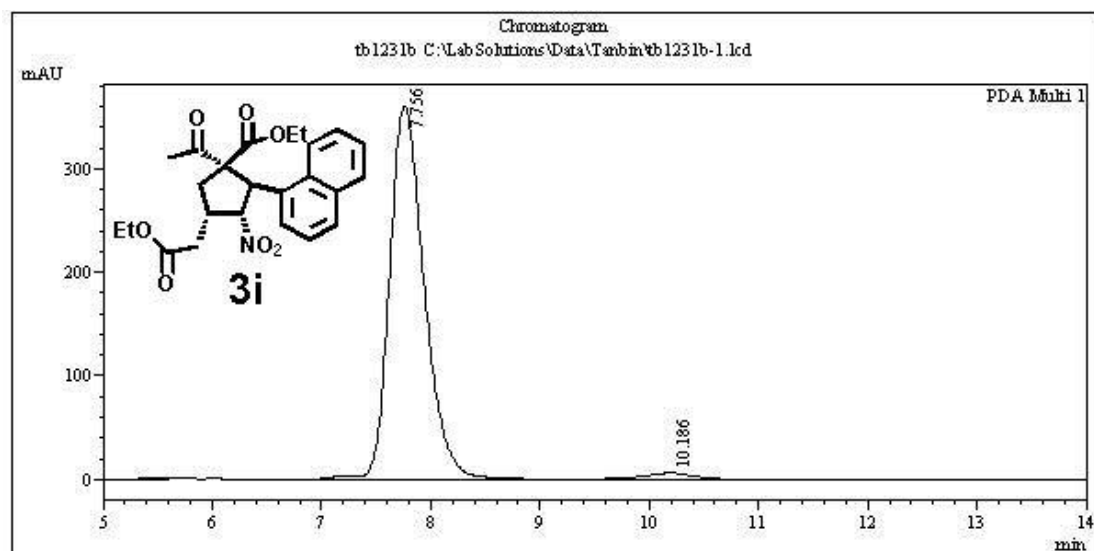
PeakTable

PDA Ch3 254nm 4nm

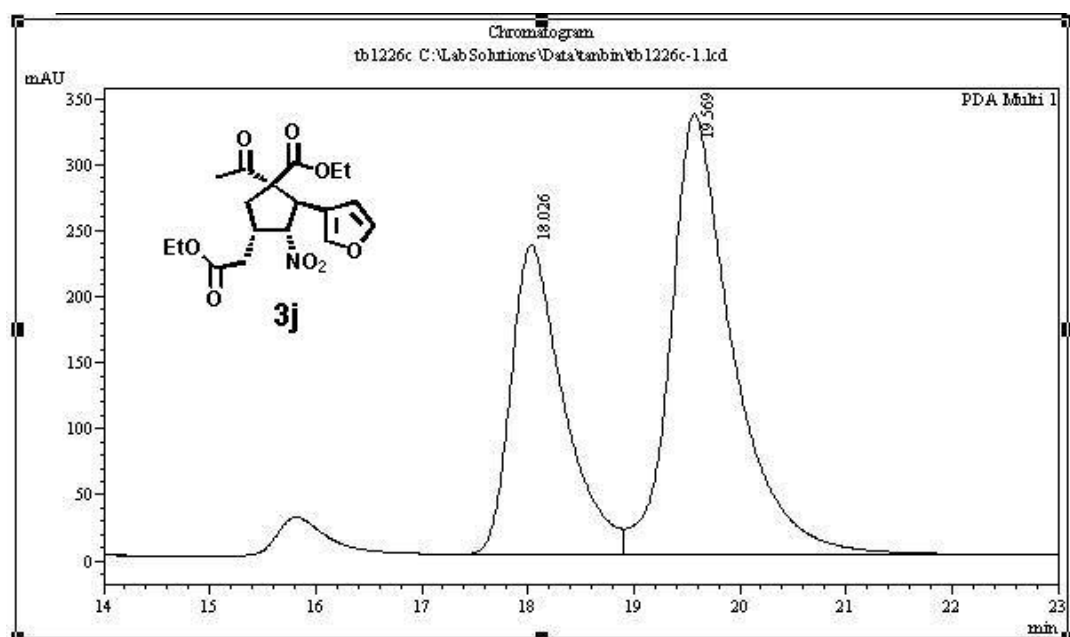
Peak#	Ret. Time	Area	Height	Area %	Height %
1	20.462	324277	7015	2.513	4.072
2	33.544	12578996	165249	97.487	95.928
Total		12903273	172263	100.000	100.000



PeakTable					
PDA Ch1 220nm 4nm					
Peak#	Ret. Time	Area	Height	Area %	Height %
1	8.014	17761993	702468	36.908	43.166
2	10.443	30363232	924894	63.092	56.834
Total		48125225	1627362	100.000	100.000



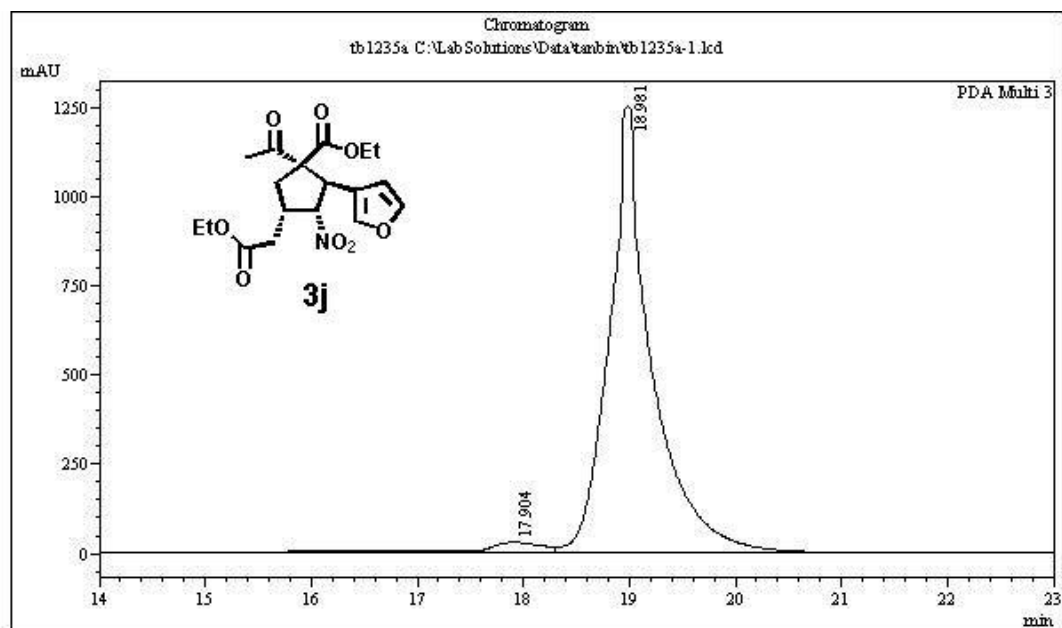
PeakTable					
PDA Ch1 220nm 4nm					
Peak#	Ret. Time	Area	Height	Area %	Height %
1	7.756	7956596	360466	97.511	98.307
2	10.186	203069	6206	2.489	1.693
Total		8159664	366672	100.000	100.000



PeakTable

PDA Ch1 210nm 4nm

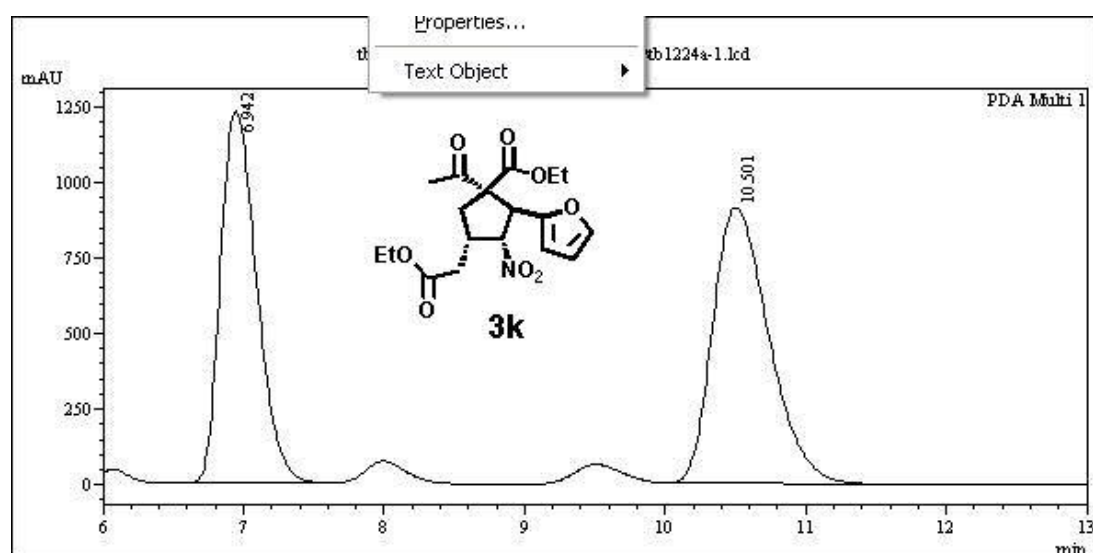
Peak#	Ret. Time	Area	Height	Area %	Height %
1	18.026	8059622	234060	37.515	41.281
2	19.569	13423997	332932	62.485	58.719
Total		21483619	566992	100.000	100.000



PeakTable

PDA Ch3 254nm 4nm

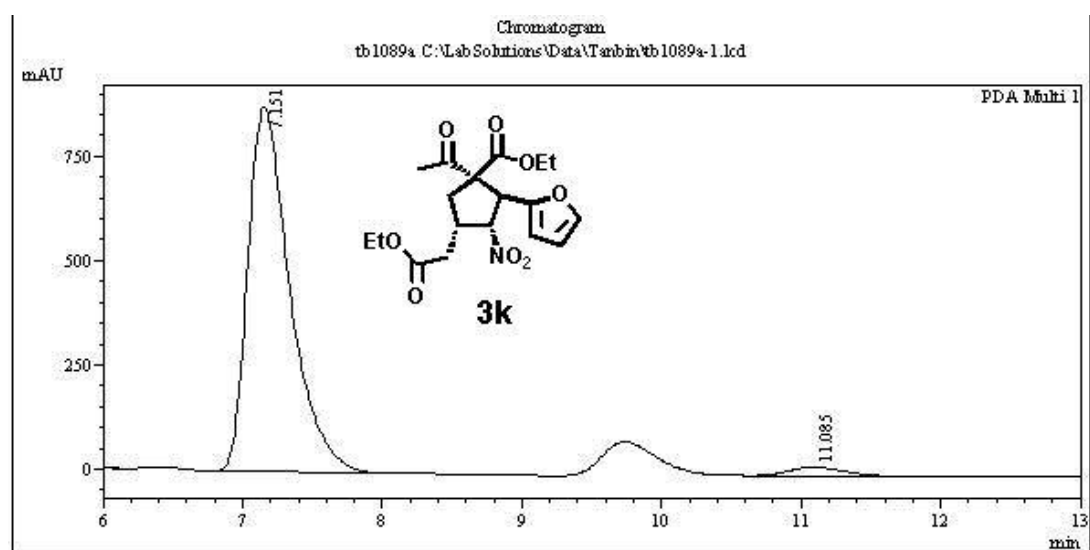
Peak#	Ret. Time	Area	Height	Area %	Height %
1	17.904	1014914	30886	2.813	2.406
2	18.981	35069428	1252908	97.187	97.594
Total		36084342	1283795	100.000	100.000



PeakTable

PDA Ch1 220nm 4nm

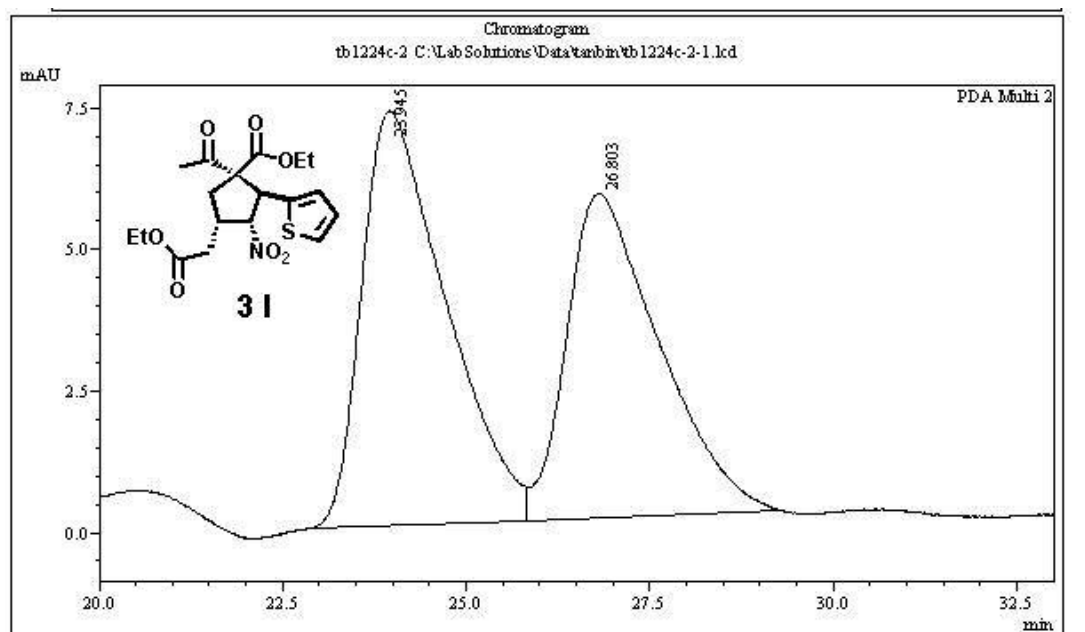
Peak#	Ret. Time	Area	Height	Area %	Height %
1	6.942	21921833	1232172	46.441	57.436
2	10.501	25282244	913126	53.559	42.564
Total		47204077	2145298	100.000	100.000



PeakTable

PDA Ch1 220nm 4nm

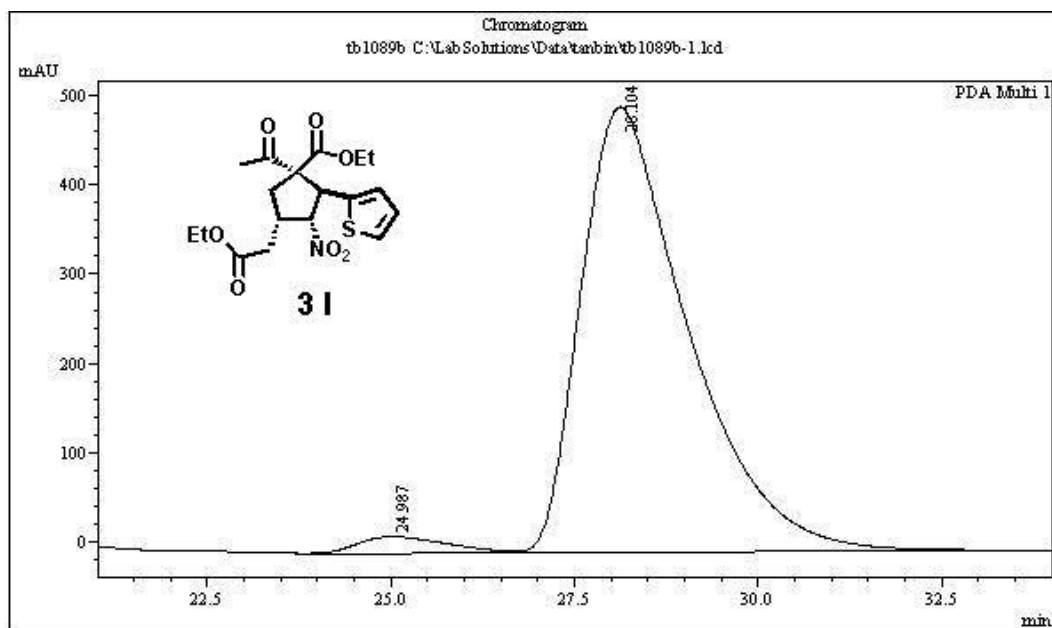
Peak#	Ret. Time	Area	Height	Area %	Height %
1	7.151	18297596	873762	96.908	97.638
2	11.085	583760	21140	3.092	2.362
Total		18881356	894903	100.000	100.000



PeakTable

PDA Ch2 230nm 4nm

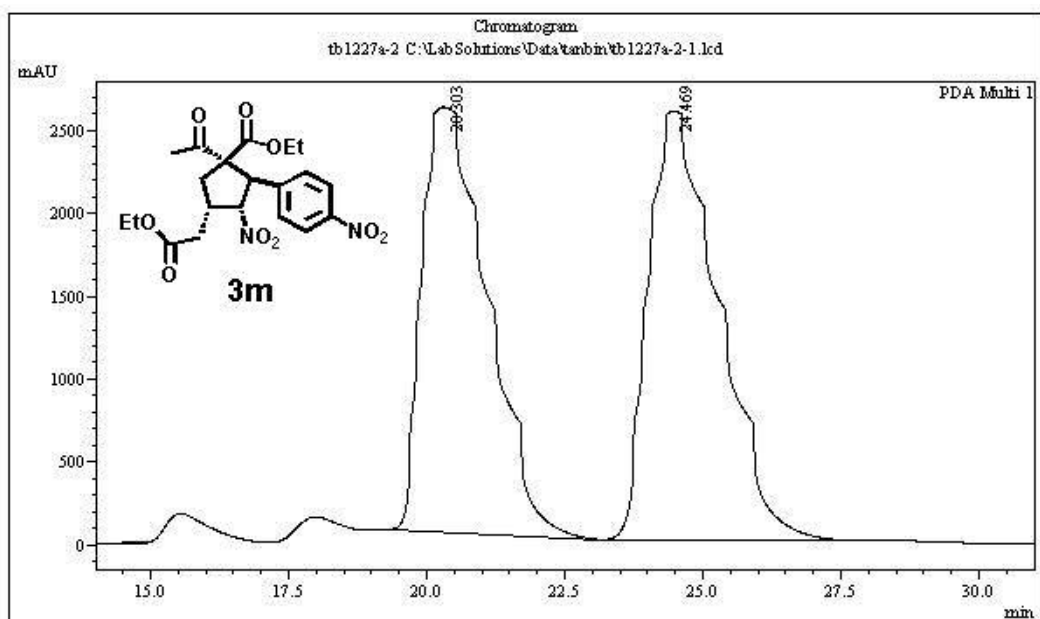
Peak#	Ret. Time	Area	Height	Area %	Height %
1	23.945	584190	7327	53.736	56.139
2	26.803	502960	5724	46.264	43.861
Total		1087151	13051	100.000	100.000



PeakTable

PDA Ch1 210nm 4nm

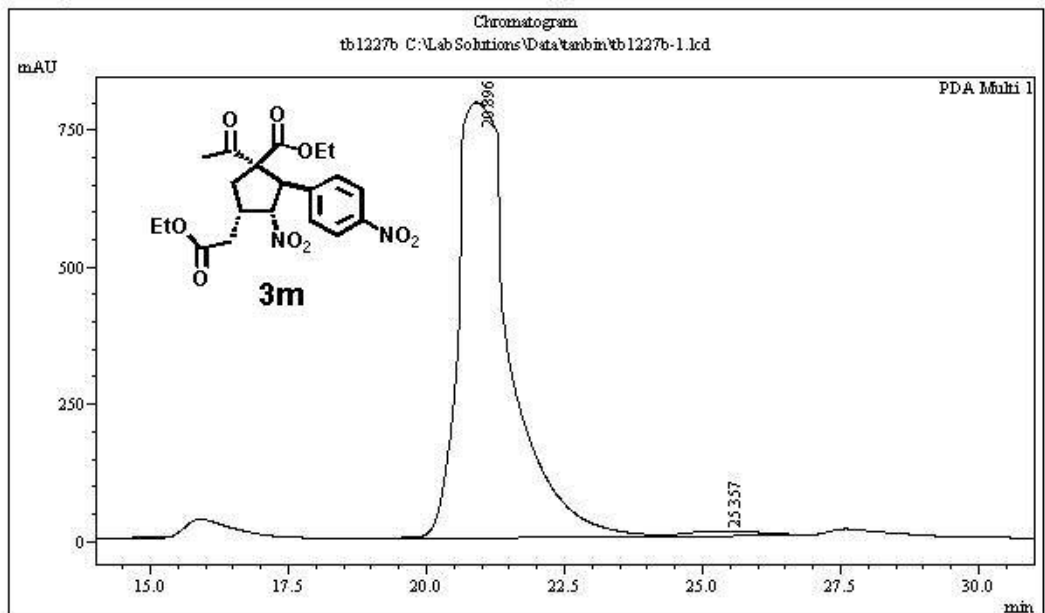
Peak#	Ret. Time	Area	Height	Area %	Height %
1	24.987	1517978	18395	2.917	3.562
2	28.104	50517843	498038	97.083	96.438
Total		52035822	516432	100.000	100.000



PeakTable

PDA Ch1 210nm 4nm

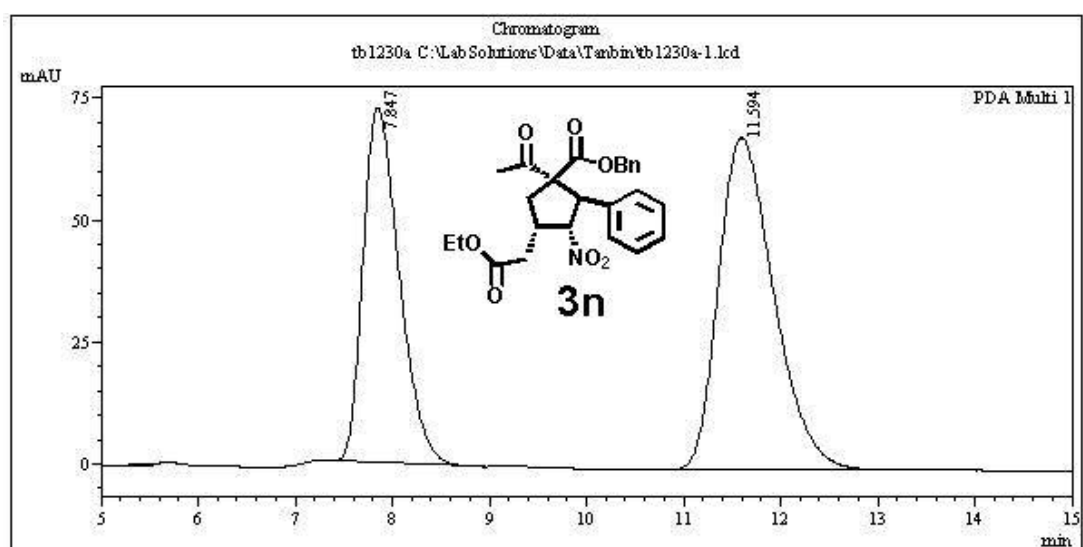
Peak#	Ret. Time	Area	Height	Area %	Height %
1	20.303	211574144	2563444	48.119	49.774
2	24.469	228118982	2586764	51.881	50.226
Total		439693126	5150209	100.000	100.000



PeakTable

PDA Ch1 210nm 4nm

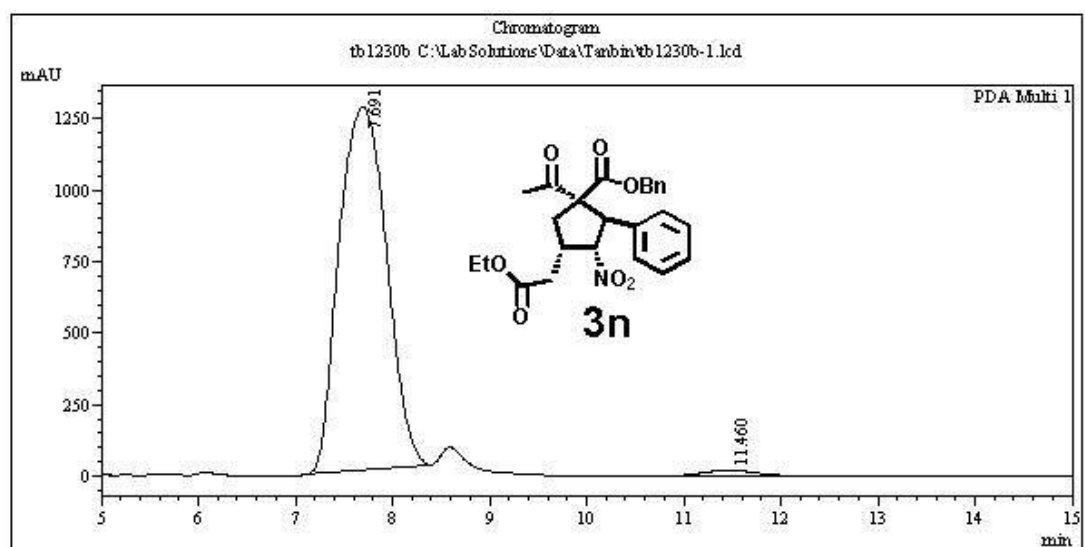
Peak#	Ret. Time	Area	Height	Area %	Height %
1	20.896	53116048	793869	98.525	98.937
2	25.357	795452	8526	1.475	1.063
Total		53911500	802395	100.000	100.000



PeakTable

PDA Ch1 220nm 4nm

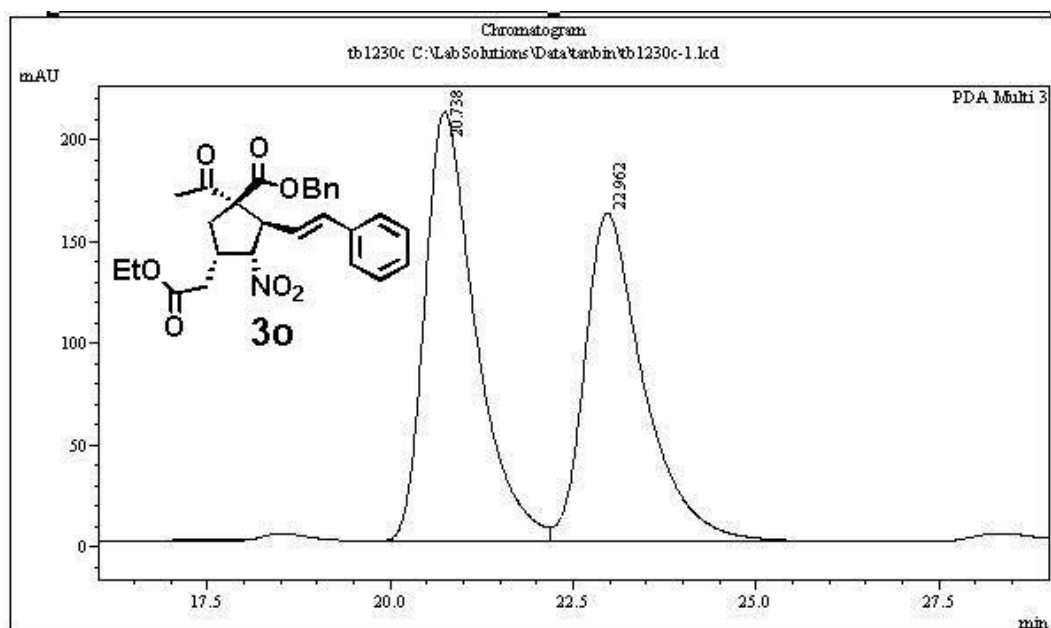
Peak#	Ret. Time	Area	Height	Area %	Height %
1	7.847	1881105	72600	41.190	51.635
2	11.594	2685778	68002	58.810	48.365
Total		4566883	140602	100.000	100.000



PeakTable

PDA Ch1 220nm 4nm

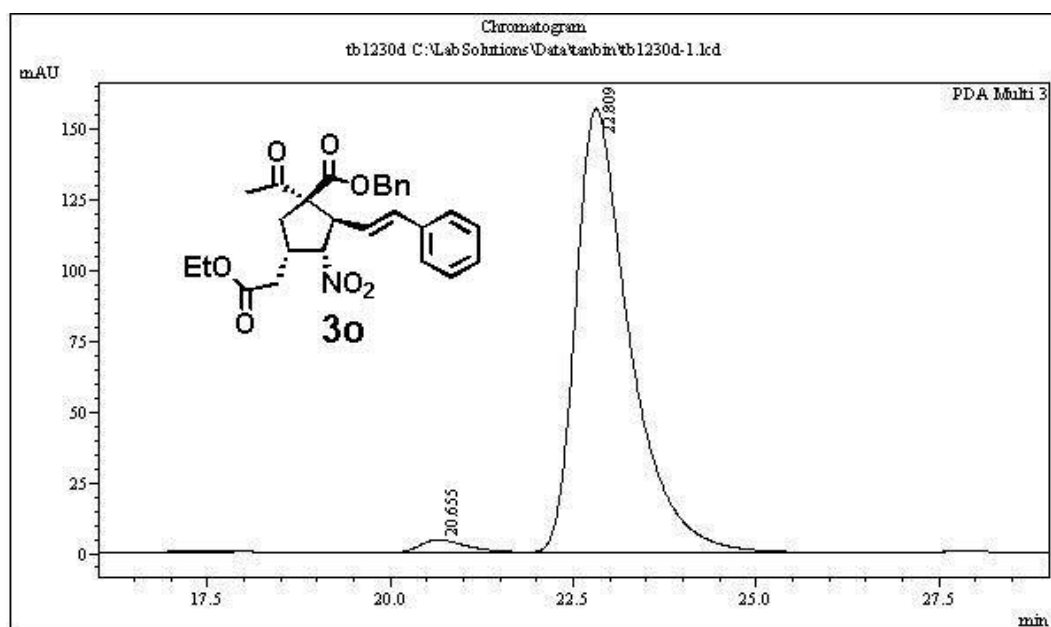
Peak#	Ret. Time	Area	Height	Area %	Height %
1	7.691	43264341	1271194	98.282	98.446
2	11.460	756084	20068	1.718	1.554
Total		44020425	1291262	100.000	100.000



PeakTable

PDA Ch3 254nm 4nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	20.738	10170695	210859	52.952	56.656
2	22.962	9036698	161313	47.048	43.344
Total		19207393	372172	100.000	100.000



PeakTable

PDA Ch3 254nm 4nm

Peak#	Ret. Time	Area	Height	Area %	Height %
1	20.655	196569	4549	2.386	2.819
2	22.809	8040203	156827	97.614	97.181
Total		8236772	161376	100.000	100.000