

Divergent synthesis of pyrrolidine and glutamic acid derivatives using macrocyclic phase-transfer catalyst under high-pressure conditions

Agata Tyszka-Gumkowska and Janusz Jurczak*

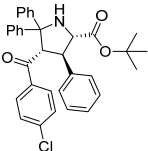
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1. Crystallographic data

The monocrystals were obtained by slowly evaporating a saturated solution of **4.2** in hexane. Single crystal X-ray diffraction measurements were carried out on a Agilent Supernova diffractometer, at 100K with graphite monochromated Cu K α radiation (1.54184 Å). The data reduction was made by using CrysAlisPRO1 software.¹ The structures were solved by direct methods and refined on F² by full-matrix least-squares by using SHELXS97 and SHELXL97.² All non-hydrogen atoms were refined as anisotropic while hydrogen atoms were placed in calculated positions, and refined in riding mode.

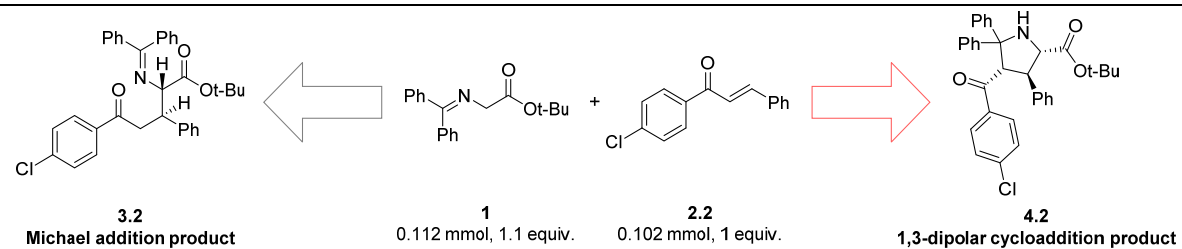
Table S1: Crystallographic data for monocrystal of compound 4.2	
compound	4.2
structure	
Empirical formula	C ₃₄ H ₃₂ Cl N O ₃
Moiety formula	C ₃₄ H ₃₂ Cl N O ₃
Formula weight	538.09
CCDC No.	2038651
Temperature	100 K
Wavelength	1.54184 Å (Cu K α)
Crystal system	triclinic
Space group	<i>P</i> -1
Unit cell dimensions	$a=9.9043(6)$ $\alpha=73.272(4)$
	$b=11.6621(6)$ $\beta=77.100(4)$
	$c=14.0011(6)$ $\gamma=68.468(5)$
Volume	1428.06(14) Å ³
<i>Z</i>	2
Density Calc.	1.390 g/cm ³
Absorption coefficient	1.457
F(000)	570.3560
Crystal	colourless block
angular range Θ	4.2 – 70.0
Index ranges	H min, max -11, 11 K min, max -14, 13. L min, max -16, 11
Reflections collected (all / independent)	5322/4857
Refinement method	Full-matrix least-squares on F^2
Absorption correction	multi-scan
Restraints / nodes / parameters	4857 / 0 / 359
Goodness-of-fit on F^2	1.0528
Final R indices [$F^2 > 2\sigma(F^2)$]	$R_1=0.0343$ $wR_2=0.0909$
R indices (all data)	$R_1=0.0373$ $wR_2=0.09036$
ρ_{\max} ρ_{\min}	0.2466 and -0.377

2. Optimization of the reaction conditions

Table S2: Investigation of optimal reaction conditions at 10 kbar pressure and 2.5 mL of MTBE as a solvent

		Michael addition product			1,3-dipolar cycloaddition product	
Base (5 equiv.)	Without catalyst	 5 mol%	Without catalyst 50 °C	 5 mol% in 50 °C	 5 mol% + 50 mol% NEt ₃	 5 mol% + 50 mol% NEt ₃ in 50 °C
CsOH·H ₂ O	W=82% d.r. 91:9	W=96% d.r. 96:4	W=90% d.r. 57:43	W=98% d.r. 92:8	W=81% d.r. 50:50	W=88% d.r. 87:13
KOH	W=85 % d.r. 88:12	W=95% d.r. 96:4	W=88% d.r. 70:30	W=95% d.r. 80:20	W=80% d.r. 80:20	W=95% d.r. 96:4
50% KOH _{aq}	W=87% d.r. 80:20	W=95% d.r. 83:17	W=61 % d.r. 96:4	W=80% d.r. 99:1	W=99% d.r. 65:35	W=94% d.r. 96:4
Cs ₂ CO ₃	W=87% d.r. 86:14	W=91% d.r. 96:4	W=72% d.r. 54:46	W=78% d.r. 82:18	W=80 % d.r. 85:15	W=99 % d.r. 99:1
K ₂ CO ₃	No reaction	No reaction	W=65% d.r. 98:2	W=54% d.r. >99:1	W=76% d.r. 97:3	W=98% d.r. 96:4
BaCO ₃	No reaction	No reaction	W=64% d.r. >99:1	W=66% d.r. >99:1	W=72% d.r. >99:1	W=93% d.r. 96:4

Table S3: Effect of different bases on the reaction course at 10 kbar pressure and 2.5 mL of MTBE as a solvent



Base (5 equiv.)	 5 mol%	 5 mol% + 50 mol% NEt₃ in 50 °C
CsOH·H ₂ O	96% d.r. 84:16	94% d.r. 96:4
KOH	96% d.r. >99:1	95% d.r. 96:4
50% KOH _{aq}	63% d.r. 80:20	94% d.r. 92:8
Cs ₂ CO ₃	W=93% d.r. 66:34	94 % d.r. 93:7
K ₂ CO ₃	No reaction	94% d.r. 96:4
BaCO ₃	No reaction	99% d.r. 98:2

Table S4: Effect of different PTC catalyst on the reaction course		
<p> $\text{Ph-CH=N(Ph)-CH}_2\text{-CO}_2\text{t-Bu}$ (1, 0.112 mmol, 1.1 equiv.) + Ph-CH=CH-CHO (2.1, 0.102 mmol, 1 equiv.) $\xrightarrow[\text{5 equiv. KOH, 2.5 mL MTBE, rt, 10 kbar, 48 h}]{\text{5 mol\% cat. PTC}}$ $\text{Ph-CH=N(Ph)-CH}_2\text{-CH(Ph)-CH}_2\text{-CO}_2\text{t-Bu}$ (3.1) </p>		
PTC catalyst	Yield	Diastereomeric ratio
	88%	92:8
	93%	95:5
	92%	95:5
	92%	95:5
	90%	97:3
	95%	96:4

Table S5: Effect of different PTC catalyst formed *in situ* on the reaction course

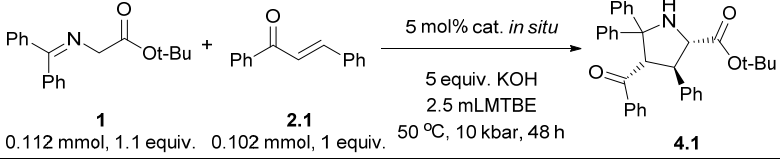
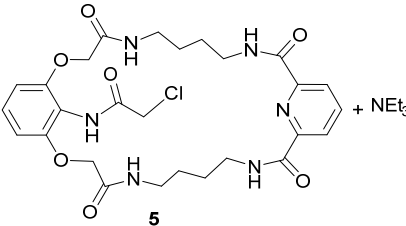
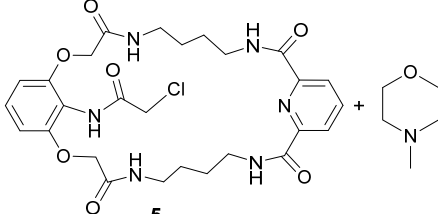
Catalyst precursor 5 mol% and 50 mol% of amine	Yield	Diastereomeric ratio
 <p>1 (0.112 mmol, 1.1 equiv.) + 2.1 (0.102 mmol, 1 equiv.)</p>	95%	96:4
 <p>5 + NE₃</p>	80%	90:10
 <p>5 + N-methylmorpholine</p>	40%	95:5

Table S6: Impact of pressure on the reaction course at 10 kbar pressure and 2.5 mL of MTBE as a solvent

 3.1 Michael addition product	 1 0.11 mmol	 2.1 0.1 mmol	 4.1 1,3-dipolar cycloaddition product
Conditions	Pressure	Yield and diastereomeric ratio	Main product
5 mol% cat. 7 , 5 equiv. KOH, room temp., 48 hours	10 kbar	95%, d.r. 94:6	3.1
	8 kbar	90%, d.r. 94:6	3.1
	5 kbar	90%, d.r. 92:8	3.1
	atmospheric pressure	52%, d.r. 69:31	3.1
5 mol% of 5 and 50 mol% of NEt ₃ , 5 equiv. KOH, 50 °C, 48 hours	10 kbar	95%, d.r. 94:6	4.1
	8 kbar	87%, d.r. 89:11	4.1
	5 kbar	82%, d.r. 63:37	4.1
	atmospheric pressure	65%, d.r. 59:41	4.1

2. Synthetic procedures and substance analysis

2.1. General remarks

All solvents and reagents were obtained from common suppliers and used as received. Flash column chromatography was performed on silica gel (230–400 mesh), thin-layer chromatography (TLC) was carried out on aluminium sheets precoated with silica gel. ^1H NMR and $^{13}\text{C}\{^1\text{H}\}$ NMR spectra were recorded on Varian 600, Varian 500 and on Bruker Mercury. Chemical shifts are reported in ppm and are set to solvent residue peak (chloroform- d_3 and acetone- d_6). The splitting pattern of multiplets is described by abbreviations (s – singlet, d – doublet, t – triplet, q – quartet, dd – doublet of doublets, m – multiplet, c – covered signal, b – broad peak). J coupling constants values are reported in Hz. Melting points were determined using a Boëtius M HMK hot-stage apparatus and were uncorrected. High resolution mass spectra (HRMS) performed with the ESI-TOF technique on a Mariner mass spectrometer from PerSeptive Biosystem.

2.2. Synthetic procedures

General Procedure A for obtaining α,β -unsaturated ketones

In accordance with the modified literature procedure.³ To a solution of KOH in water (8 g of KOH in 8 mL of water) the solution of appropriate methyl ketone (1.0 equiv., 5.0 mmol) and aldehyde (1.0 equiv, 5.0 mmol) in MeOH (10 mL) at 0 °C was added dropwise. Typically, the mixture started to solidified during addition of substrates. The reaction was monitored by TLC. After completion, H₂O (150 mL) was added at 0 °C and the resulting precipitate filtered. Recrystallization from EtOH afforded the pure α,β -unsaturated ketones. In the cases where precipitation did not occur, the reaction mixture was extracted with DCM (3 x 50 mL). Organic layers were combined, washed with brine, dried over Na₂SO₄, filtered and evaporated to dryness in vacuo. Subsequent purification by flash column chromatography using hexane:ethyl acetate (95:5→9:1) as an eluent afforded the pure products.

General Procedure B for obtaining α,β -unsaturated ketones

In accordance with the modified literature procedure.⁴ To a solution of the appropriate methyl ketone (1.0 equiv., 5.0 mmol) and aldehyde (1.0 equiv., 5.0 mmol) in MeOH (10 mL) at 0 °C was added a 10% aqueous solution of KOH (30 mL, dropwise). Typically, the mixture started to solidified after 15 minutes. The reaction was monitored by TLC. After completion of the reaction, H₂O (150 mL) was added at 0 °C and the resulting precipitate was filtered. Recrystallization from EtOH afforded the pure α,β -unsaturated ketones. In the cases where precipitation did not occur, the reaction mixture was extracted with DCM (3 x 50 mL). Organic

layers were combined, washed with brine, dried over Na₂SO₄, filtered and evaporated to dryness in vacuo. Subsequent purification by flash column chromatography using hexane:ethyl acetate (95:5→9:1) as an eluent afforded the pure products.

General Procedure C for obtaining Michael adducts **3.2-3.23**

Glycine ketamine **1** (33 mg, 0.112 mmol, 1.1 equiv.), appropriate α , β -unsaturated ketone (1 equiv., 0.102 mmol), PTC **7** catalyst (3.5 mg, 5 mol%, 0.0051 mmol) and KOH (5 equiv., 28.5 mg, 0.510 mmol) were placed into a 2.5 mL Teflon reaction vessel. Then the reaction vessel was filled with MTBE and screwed tightly to avoid any air bubbles inside. The reaction was carried out in a high-pressure apparatus for 48 hours, using a pressure of 10kbar. After complexation, the reaction mixture was filtered through a pad of silica gel and magnesium sulphate, washing the product with DCM. After concentration, the crude product was purified by preparative HPLC (loading as a solution in DCM), eluting with a solvent system of 5% AcOEt + 1% NEt₃ in hexane to separate the excess of ketamine **1**. Combined fractions were concentrated and dried to receive the final compound.

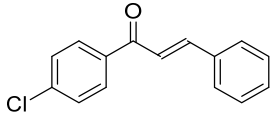
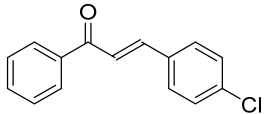
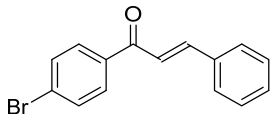
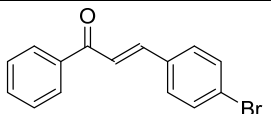
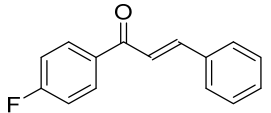
In experiments on larger scale glycine ketamine **1** (330 mg, 1.12 mmol, 1.1 equiv.), α , β -unsaturated ketone **2.1** (212 mg, 1.02 mmol, 1 equiv.), PTC **7** catalyst (35 mg, 5 mol%, 0.051 mmol) and KOH (5 equiv., 285 mg, 5.10 mmol) were placed into a 25 mL Teflon reaction vessel. After purification product were obtained as a clear oil (467 mg, 91%, d.r. 90:10).

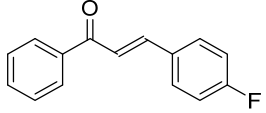
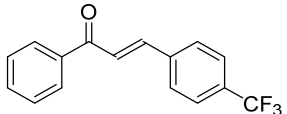
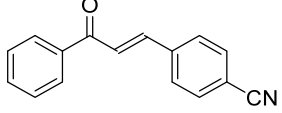
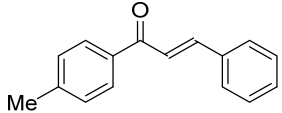
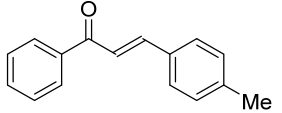
General Procedure D for obtaining 1,3-dipolar cycloaddition products **4.2-4.21**

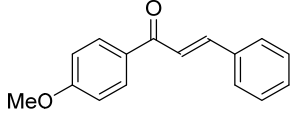
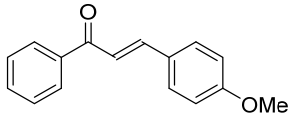
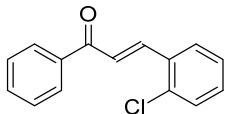
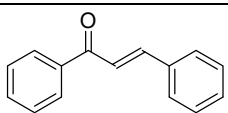
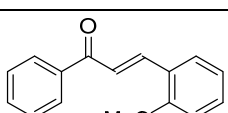
Glycine ketamine **1** (33 mg, 0.112 mmol, 1.1 equiv.), appropriate α , β -unsaturated ketone (1 equiv., 0.102 mmol), macrocyclic precursor **5** (3 mg, 5 mol%, 0.0051 mmol), NEt₃ (50 μ L, 5 ekw.) and KOH (5 equiv., 28.5 mg, 0.510 mmol) were placed into a 2.5 mL Teflon reaction vessel. Then the reaction vessel was filled with MTBE and screwed tightly to avoid any air bubbles inside. The reaction was carried out in a high-pressure apparatus for 48 hours, using a pressure of 10kbar and temperature 50 °C. After complexation, the reaction mixture was filtered through a pad of silica gel and magnesium sulphate, washing the product with DCM. After concentration, the crude product was purified by preparative HPLC (loading as a solution in DCM), eluting with a solvent system of 5% AcOEt + 1% NEt₃ in hexane to separate the excess of ketamine **1**. Combined fractions were concentrated and dried to receive the final compound.

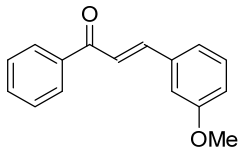
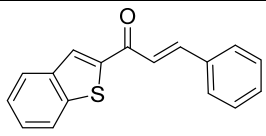
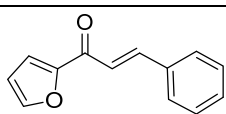
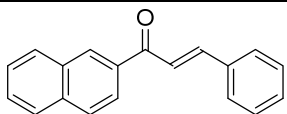
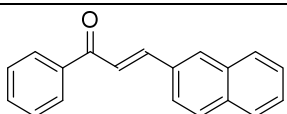
In experiments on larger scale glycine ketamine **1** (330 mg, 1.12 mmol, 1.1 equiv.), α , β -unsaturated ketone **2.1** (212 mg, 1.02 mmol, 1 equiv.), macrocyclic precursor **5** (30 mg, 5 mol%, 0.051 mmol), NEt₃ (500 μ L, 5 ekw.) and KOH (5 equiv., 285 mg, 5.10 mmol) were placed into a 25 mL Teflon reaction vessel. After purification product were obtained as a clear oil (410 mg, 80%, d.r. 92:8).

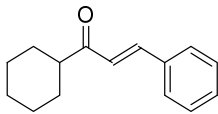
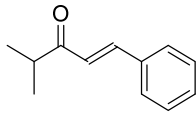
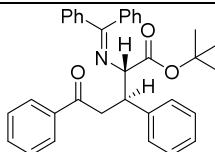
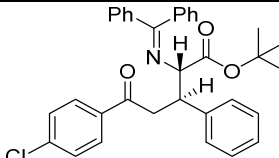
2.3. Substance analysis

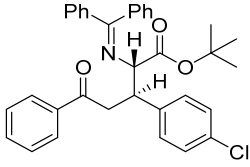
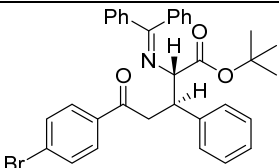
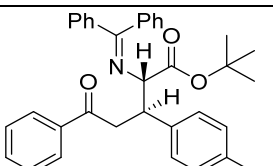
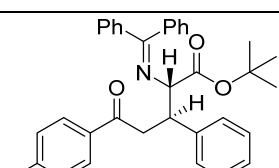
	<p>Following Procedure A and using 4-chloroacetophenone (776 mg, 5 mmoli) and benzaldehyde (530 mg, 5mmoli) as starting materials, compound 2.2 was obtained as yellowish crystals (1068 mg, 88%).^{5,8}</p> <p>¹H NMR (400 MHz, CDCl₃) δ 8.00 – 7.93 (m, 2H), 7.81 (d, <i>J</i> = 15.7 Hz, 1H), 7.68 – 7.60 (m, 2H), 7.54 – 7.37 (m, 6H). ¹³C NMR (101 MHz, CDCl₃) δ 189.1, 145.3, 139.2, 136.5, 134.7, 130.7, 129.9, 129.0, 128.9, 128.5, 121.6.</p>
	<p>Following Procedure A and using acetophenone (601 mg, 5 mmol) and 4-chlorobenzaldehyde (703 mg, 5mmoli) as starting materials, compound 2.3 was obtained as yellowish crystals (1043 mg, 86%).^{3,7}</p> <p>¹H NMR (400 MHz, CDCl₃) δ 8.04 – 7.97 (m, 1H), 7.76 (d, <i>J</i> = 15.7 Hz, 1H), 7.63 – 7.54 (m, 2H), 7.54 – 7.45 (m, 2H), 7.39 (d, <i>J</i> = 8.5 Hz, 1H). ¹³C NMR (101 MHz, CDCl₃) δ 190.2, 143.3, 138.1, 136.4, 133.4, 132.9, 129.6, 129.2, 128.7, 128.5, 122.5.</p>
	<p>Following Procedure A and using 4-bromoacetophenone (995 mg, 5 mmol) and benzaldehyde (530 mg, 5mmoli) as starting materials, compound 2.4 was obtained as yellowish crystals (1307 mg, 92%).^{6,7}</p> <p>¹H NMR (500 MHz, cdcl₃) δ 7.83 – 7.77 (m, 2H), 7.73 (d, <i>J</i> = 15.7 Hz, 1H), 7.58 – 7.52 (m, 4H), 7.39 (d, <i>J</i> = 15.7 Hz, 1H), 7.36 – 7.31 (m, 3H). ¹³C NMR (126 MHz, cdcl₃) δ 189.3, 145.4, 136.9, 134.7, 131.9, 130.7, 130.0, 128.9, 128.5, 127.9, 121.5.</p>
	<p>Following Procedure A and using acetophenone (601 mg, 5 mmol) and 4-bromobenzaldehyde (925 mg, 5mmoli) as starting materials, compound 2.5 was obtained as yellowish crystals (1278 mg, 89%).^{3,7}</p> <p>¹H NMR (400 MHz, CDCl₃) δ 8.01 (d, <i>J</i> = 5.3, 3.4 Hz, 1H), 7.74 (d, <i>J</i> = 15.7 Hz, 1H), 7.63 – 7.46 (m, 4H). ¹³C NMR (101 MHz, CDCl₃) δ 190.2, 143.3, 138.0, 133.8, 132.9, 132.2, 129.8, 128.7, 128.5, 124.8, 122.6.</p>
	<p>Following Procedure A and using 4-fluoroacetophenone (776 mg, 5 mmol) and benzaldehyde (530 mg, 5mmoli) as starting materials, compound 2.6 was obtained as yellowish crystals (1018 mg, 90%).^{7,8}</p> <p>¹H NMR (400 MHz, CDCl₃) δ 8.05 (dd, <i>J</i> = 7.6, 5.5 Hz, 2H), 7.81 (d, <i>J</i> = 15.7 Hz, 1H), 7.64 (d, <i>J</i> = 2.9 Hz, 2H), 7.59 – 7.34 (m, 4H), 7.19 (dd, <i>J</i> = 24.6, 16.8 Hz, 2H). ¹³C NMR (101 MHz, CDCl₃) δ 188.80, 165.6 (d, <i>J</i>_{C-F} = 252.9 Hz), 145.0, 134.8, 134.6 (d, <i>J</i>_{C-F} = 2.8 Hz), 131.1 (d, <i>J</i>_{C-F} = 9.1 Hz), 130.6, 128.9, 128.5, 121.6, 115.7 (d, <i>J</i>_{C-F} = 22.0 Hz).</p>

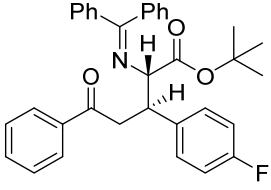
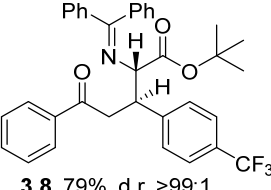
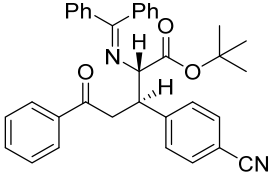
	<p>Following Procedure A and using acetophenone (601 mg, 5 mmol) and 4-fluorobenzaldehyde (620mg, 5mmoli) as starting materials, compound 2.7 was obtained as yellowish crystals (1052 mg, 93%).⁸</p> <p>¹H NMR (400 MHz, CDCl₃) δ 8.01 (d, <i>J</i> = 7.4 Hz, 2H), 7.77 (d, <i>J</i> = 15.7 Hz, 1H), 7.67 – 7.54 (m, 3H), 7.54 – 7.38 (m, 3H), 7.10 (t, <i>J</i> = 8.5 Hz, 2H). ¹³C NMR (101 MHz, CDCl₃) δ 190.3, 164.1 (d, <i>J</i>_{C-F} = 251.9 Hz), 143.5, 138.2, 132.8, 131.2 (d, <i>J</i>_{C-F} = 3.3 Hz), 130.4, 130.3, 128.6, 128.5, 121.8 (d, <i>J</i>_{C-F} = 2.3 Hz), 116.1 (d, <i>J</i>_{C-F} = 21.9 Hz).</p>
	<p>Following Procedure B and using acetophenone (601 mg, 5 mmol) and 4-(trifluoromethyl)benzaldehyde (871 mg, 5mmoli) as starting materials, compound 2.8 was obtained as yellow flakes (870 mg, 63%).⁸</p> <p>¹H NMR (400 MHz, CDCl₃) δ 8.03 (d, <i>J</i> = 7.1 Hz, 2H), 7.90 – 7.40 (m, 9H). ¹³C NMR (101 MHz, CDCl₃) δ 190.0, 142.7, 138.3, 137.8, 133.1, 128.7, 128.5, 128.5, 126.4, 125.4, 125.2, 124.3.</p>
	<p>Following Procedure B and using acetophenone (601 mg, 5 mmol) and 4-cyanobenzaldehyde (606 mg, 5mmoli) as starting materials, compound 2.9 was obtained as yellowish crystals (851 mg, 73%).³</p> <p>¹H NMR (400 MHz, CDCl₃) δ 8.02 (d, <i>J</i> = 7.0 Hz, 2H), 7.65 (m, 9H). ¹³C NMR (101 MHz, CDCl₃) δ 189.7, 142.0, 139.2, 137.7, 133.3, 132.7, 125.1, 118.3, 113.5, 29.7.</p>
	<p>Following Procedure A and using 4-methylacetophenone (671 mg, 5 mmol) and benzaldehyde (530 mg, 5mmoli) as starting materials, compound 2.10 was obtained as yellowish crystals (856 mg, 77%).^{5,8}</p> <p>¹H NMR (400 MHz, CDCl₃) δ 7.94 (d, <i>J</i> = 8.1 Hz, 2H), 7.81 (d, <i>J</i> = 15.7 Hz, 1H), 7.63 (dd, <i>J</i> = 6.4, 2.8 Hz, 2H), 7.53 (d, <i>J</i> = 15.7 Hz, 1H), 7.41 (dd, <i>J</i> = 4.9, 1.6 Hz, 3H), 7.30 (d, <i>J</i> = 8.0 Hz, 2H), 2.43 (s, 3H). ¹³C NMR (101 MHz, CDCl₃) δ 189.9, 144.4, 143.6, 135.7, 135.0, 130.4, 129.3, 128.9, 128.7, 128.4, 122.2, 21.7.</p>
	<p>Following Procedure A and using acetophenone (601 mg, 5 mmol) and 4-methylbenzaldehyde (606 mg, 5mmoli) as starting materials, compound 2.11 was obtained as yellowish crystals (889 mg, 80%).⁸</p> <p>¹H NMR (400 MHz, CDCl₃) δ 8.05 – 7.98 (m, 2H), 7.79 (d, <i>J</i> = 15.7 Hz, 1H), 7.62 – 7.45 (m, 6H), 7.23 (d, <i>J</i> = 8.0 Hz, 2H), 2.39 (s, 3H). ¹³C NMR (101 MHz, CDCl₃) δ 190.6, 144.9, 141.1, 138.4, 132.6, 132.2, 129.7, 128.6, 128.5, 128.5, 121.2, 21.5.</p>

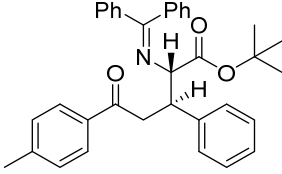
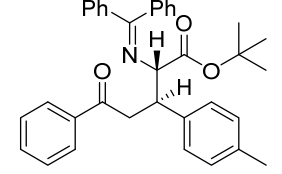
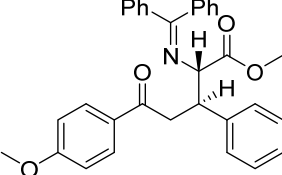
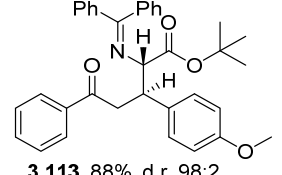
	<p>Following Procedure A and using 4-methoxyacetophenone (751 mg, 5 mmol) and benzaldehyde (530 mg, 5mmoli) as starting materials, compound 2.12 was obtained as yellowish crystals (990 mg, 83%).^{5,6,7}</p> <p>¹H NMR (400 MHz, CDCl₃) δ 8.04 (d, <i>J</i> = 8.8 Hz, 2H), 7.80 (d, <i>J</i> = 15.7 Hz, 1H), 7.69 – 7.60 (m, 2H), 7.54 (d, <i>J</i> = 15.7 Hz, 1H), 7.47 – 7.33 (m, 3H), 6.99 (d, <i>J</i> = 8.8 Hz, 2H), 3.89 (s, <i>J</i> = 11.0 Hz, 3H). ¹³C NMR (101 MHz, CDCl₃) δ 188.7, 163.5, 143.9, 135.1, 131.1, 130.8, 130.3, 128.9, 128.4, 121.9, 113.9, 55.5.</p>
	<p>Following Procedure A and using acetophenone (601 mg, 5 mmol) and 4-methoxybenzaldehyde (681 mg, 5mmoli) as starting materials, compound 2.13 was obtained as yellowish crystals (1132 mg, 95%).^{7,8}</p> <p>¹H NMR (400 MHz, CDCl₃) δ 8.01 (d, <i>J</i> = 7.1 Hz, 2H), 7.79 (d, <i>J</i> = 15.7 Hz, 1H), 7.58 (dd, <i>J</i> = 16.2, 8.0 Hz, 3H), 7.50 (t, <i>J</i> = 7.4 Hz, 2H), 7.41 (d, <i>J</i> = 15.6 Hz, 1H), 6.94 (d, <i>J</i> = 8.6 Hz, 2H), 3.86 (s, 3H). ¹³C NMR (101 MHz, CDCl₃) δ 190.6, 161.7, 144.7, 138.5, 132.5, 130.2, 128.6, 128.4, 127.7, 119.9, 114.4, 55.4.</p>
	<p>Following Procedure B and using acetophenone (601 mg, 5 mmol) and 2-chlorobenzaldehyde (703 mg, 5mmoli) as starting materials, compound 2.14 was obtained as yellowish crystals (364 mg, 30%).⁹</p> <p>¹H NMR (400 MHz, CDCl₃) δ 8.18 (d, <i>J</i> = 15.8 Hz, 1H), 8.01 (dd, <i>J</i> = 5.2, 3.3 Hz, 2H), 7.75 (dd, <i>J</i> = 6.9, 2.5 Hz, 1H), 7.62 – 7.56 (m, 1H), 7.55 – 7.41 (m, 4H), 7.36 – 7.28 (m, 2H). ¹³C NMR (101 MHz, CDCl₃) δ 190.5, 140.6, 137.9, 135.5, 133.3, 132.9, 131.1, 130.3, 128.6, 128.6, 127.8, 127.1, 124.9.</p>
	<p>Following Procedure B and using acetophenone (601 mg, 5 mmol) and 3-chlorobenzaldehyde (703 mg, 5mmoli) as starting materials, compound 2.15 was obtained as yellowish crystals (813 mg, 67%).⁹</p> <p>¹H NMR (500 MHz, cdcl₃) δ 7.98 – 7.90 (m, 2H), 7.65 (d, <i>J</i> = 15.7 Hz, 1H), 7.57 – 7.49 (m, 2H), 7.45 (d, <i>J</i> = 7.2 Hz, 1H), 7.44 – 7.39 (m, 3H), 7.32 – 7.26 (m, 2H). ¹³C NMR (126 MHz, cdcl₃) δ 190.1, 143.0, 137.9, 136.7, 134.9, 133.0, 130.3, 130.2, 128.7, 128.5, 127.9, 126.8, 123.2.</p>
	<p>Following Procedure B and using acetophenone (601 mg, 5 mmol) and 2-methoxybenzaldehyde (681 mg, 5mmoli) as starting materials, compound 2.16 was obtained as yellow solid (870 mg, 73%).⁸</p> <p>¹H NMR (400 MHz, CDCl₃) δ 8.12 (d, <i>J</i> = 15.9 Hz, 1H), 8.05 – 7.97 (m, 2H), 7.66 – 7.53 (m, 3H), 7.52 – 7.46 (m, 2H), 7.38 (ddd, <i>J</i> = 8.3, 7.4, 1.7 Hz, 1H), 6.99 (td, <i>J</i> = 7.5, 1.1 Hz, 1H), 6.94 (dd, <i>J</i> = 8.3, 1.1 Hz, 1H), 3.91 (s, 3H).</p>

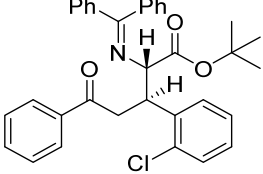
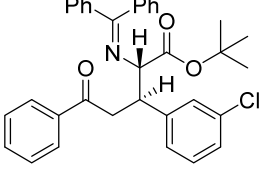
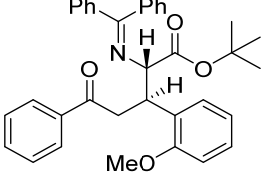
	¹³ C NMR (101 MHz, CDCl ₃) δ 191.1, 158.8, 140.4, 138.6, 132.5, 131.7, 129.2, 128.5, 123.9, 122.9, 120.8, 111.3, 55.6.
	Following Procedure B and using acetophenone (601 mg, 5 mmol) and 3-methoxybenzaldehyde (681 mg, 5mmoli) as starting materials, compound 2.17 was obtained as yellow solid (727 mg, 61%). ⁸ ¹ H NMR (400 MHz, CDCl ₃) δ 8.01 (dd, <i>J</i> = 8.4, 1.4 Hz, 2H), 7.77 (d, <i>J</i> = 15.7 Hz, 1H), 7.62 – 7.55 (m, 1H), 7.54 – 7.44 (m, 3H), 7.33 (t, <i>J</i> = 7.9 Hz, 1H), 7.27 – 7.21 (m, 1H), 7.15 (dd, <i>J</i> = 2.6, 1.6 Hz, 1H), 6.96 (ddd, <i>J</i> = 8.2, 2.6, 1.0 Hz, 1H), 3.85 (s, 3H). ¹³ C NMR (101 MHz, CDCl ₃) δ 190.5, 159.9, 144.7, 138.2, 136.3, 132.8, 129.9, 128.6, 128.5, 122.5, 121.1, 116.3, 113.5, 55.4.
	Following Procedure B and using 2-acetylbenzothiophene (882 mg, 5 mmol) and benzaldehyde (530 mg, 5 mmol) as starting materials, compound 2.18 was obtained as yellowish crystals (950 mg, 72%). ¹⁰ ¹ H NMR (500 MHz, CDCl ₃) δ 8.12 (s, 1H), 7.98 – 7.84 (m, 3H), 7.69 (dd, <i>J</i> = 6.6, 2.7 Hz, 2H), 7.55 (d, <i>J</i> = 15.6 Hz, 1H), 7.51 – 7.38 (m, 5H). ¹³ C NMR (126 MHz, CDCl ₃) δ 183.4, 145.2, 144.4, 142.7, 139.3, 134.7, 130.7, 129.0, 128.8, 128.6, 127.4, 125.9, 125.0, 123.0, 121.2.
	Following Procedure B and using 2-acetylfurane (550 mg, 5 mmol) and benzaldehyde (530 mg, 5mmoli) as starting materials, compound 2.19 was obtained as beige crystals (922 mg, 93%). ⁸ ¹ H NMR (400 MHz, CDCl ₃) δ 7.88 (d, <i>J</i> = 15.8 Hz, 1H), 7.65 (dt, <i>J</i> = 4.1, 2.7 Hz, 3H), 7.45 (d, <i>J</i> = 15.8 Hz, 1H), 7.42 – 7.39 (m, 3H), 7.33 (dd, <i>J</i> = 3.7, 0.8 Hz, 1H), 6.59 (dd, <i>J</i> = 3.5, 1.7 Hz, 1H). ¹³ C NMR (101 MHz, CDCl ₃) δ 178.0, 154.4, 146.5, 143.9, 134.8, 130.6, 128.9, 128.5, 121.2, 117.5, 112.5.
	Following Procedure A and using 2-acetylnaphthalene (851 mg, 5 mmol) and benzaldehyde (530 mg, 5mmoli) as starting materials, compound 2.20 was obtained as yellowish crystals (1175mg, 91%). ^{5,8} ¹ H NMR (400 MHz, CDCl ₃) δ 8.54 (s, 1H), 8.11 (d, <i>J</i> = 7.8 Hz, 1H), 7.99 (d, <i>J</i> = 6.9 Hz, 1H), 7.98 – 7.83 (m, 3H), 7.78 – 7.66 (m, 3H), 7.60 (dd, <i>J</i> = 15.4, 7.5 Hz, 2H), 7.44 (d, <i>J</i> = 2.3 Hz, 3H). ¹³ C NMR (101 MHz, CDCl ₃) δ 190.3, 144.8, 135.6, 135.5, 135.0, 132.6, 130.5, 129.9, 129.5, 129.0, 128.6, 128.5, 128.4, 127.8, 126.8, 124.5, 122.2.
	Following Procedure B and using acetophenone (601 mg, 5 mmol) and 2-naphthaldehyde (776 mg, 5mmoli) as starting materials, compound 2.21 was obtained as yellowish crystals (1150 mg, 89%). ^{4,9} ¹ H NMR (400 MHz, CDCl ₃) δ 8.05 (dd, <i>J</i> = 6.8, 5.3 Hz, 3H), 7.98 (d, <i>J</i> = 15.7 Hz, 1H), 7.87 (dt, <i>J</i> = 14.8, 5.2 Hz, 3H), 7.80 (dd, <i>J</i> = 8.6, 1.5 Hz, 1H), 7.62 (dd, <i>J</i> = 21.1, 11.6 Hz, 2H), 7.52 (dd, <i>J</i> = 8.6, 4.7 Hz, 4H). ¹³ C NMR (101

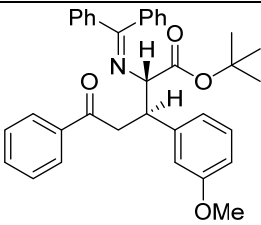
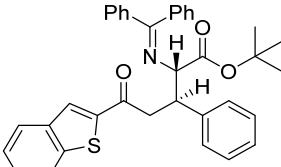
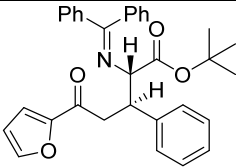
	MHz, CDCl ₃) δ 190.5, 144.9, 143.6, 138.3, 134.4, 133.4, 132.8, 132.4, 130.6, 128.7, 128.6, 128.5, 127.8, 127.4, 126.8, 123.7, 122.3.
	Following Procedure B and using 1-cyclohexylethanone (631 mg, 5 mmol) and benzaldehyde (530 mg, 5mmoli) as starting materials, compound 2.22 was obtained as white flakes (997 mg, 93%). ⁶ ¹H NMR (400 MHz, CDCl ₃) δ 7.63 – 7.49 (m, 3H), 7.42 – 7.30 (m, 3H), 6.81 (d, <i>J</i> = 16.0 Hz, 1H), 2.65 (tt, <i>J</i> = 11.3, 3.3 Hz, 1H), 1.95 – 1.62 (m, 5H), 1.51 – 1.16 (m, 5H). ¹³C NMR (101 MHz, CDCl ₃) δ 203.1, 142.2, 134.8, 130.3, 128.9, 128.2, 124.7, 49.4, 28.7, 25.9, 25.8.
	Following Procedure B and using 3-methyl-2-butanone (430 mg, 5 mmol) and benzaldehyde (530 mg, 5mmoli) as starting materials, compound 2.23 was obtained as yellow oil (261mg, 30%). ¹¹ ¹H NMR (400 MHz, CDCl ₃) δ 7.61 (d, <i>J</i> = 16.0 Hz, 1H), 7.58 – 7.53 (m, 1H), 7.38 (dd, <i>J</i> = 4.2, 2.4 Hz, 3H), 6.82 (d, <i>J</i> = 16.0 Hz, 1H), 2.93 (dt, <i>J</i> = 13.8, 6.9 Hz, 1H), 1.18 (d, <i>J</i> = 6.9 Hz, 7H). ¹³C NMR (101 MHz, CDCl ₃) δ 203.8, 142.4, 134.7, 130.3, 128.9, 128.3, 124.5, 39.3, 18.5.
 3.1 , 95%, d.r. 94:6	Following Procedure C using α,β-unsaturated ketone 2.1 (21 mg, 0.102 mmol) as starting material, compound 3.1 was obtained as clear oil (49 mg, 95%). ^{12,13,14,15} ¹H NMR (400 MHz, Acetone) δ 7.97 (dd, <i>J</i> = 5.2, 3.3 Hz, 2H), 7.71 – 7.65 (m, 2H), 7.61 – 7.55 (m, 1H), 7.51 – 7.35 (m, 8H), 7.23 – 7.09 (m, 5H), 6.88 (dd, <i>J</i> = 7.0, 2.3 Hz, 2H), 4.26 – 4.16 (m, 2H), 3.79 – 3.58 (m, 2H), 1.28 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 198.9, 171.9, 170.5, 142.9, 140.6, 138.5, 137.5, 133.8, 131.5, 129.8, 129.7, 129.6, 129.6, 129.4, 129.2, 129.06, 129.05, 128.7, 127.5, 81.6, 72.2, 46.1, 41.2, 28.2. HPLC : Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 11.48 min, 15.08 min.
 3.2 , 96%, d.r. >99:1	Following Procedure C using α,β-unsaturated ketone 2.2 (24.5 mg, 0.102 mmol) as starting material, compound 3.2 was obtained as clear foam (52 mg, 96%, mp 129.6 °C). ^{12,14,15} ¹H NMR (500 MHz, acetone) δ 7.98 (d, <i>J</i> = 8.4 Hz, 1H), 7.69 (d, <i>J</i> = 8.5 Hz, 1H), 7.57 (dt, <i>J</i> = 13.1, 6.8 Hz, 2H), 7.51 – 7.42 (m, 6H), 7.38 (t, <i>J</i> = 7.4 Hz, 2H), 7.23 (d, <i>J</i> = 3.4 Hz, 3H), 6.99 – 6.90 (m, 1H), 4.25 – 4.17 (m, 1H), 3.80 – 3.71 (m, 1H), 3.65 (dd, <i>J</i> = 17.1, 2.9 Hz, 1H), 1.29 (s, 9H). ¹³C NMR (126 MHz, acetone) δ 197.6, 171.0, 169.2, 140.8, 139.4, 137.2, 136.2, 132.8, 132.4, 131.8, 130.5, 129.7, 128.6, 128.5, 128.4, 128.1, 127.9, 127.9, 127.5, 80.7, 70.8, 44.4, 40.0, 27.1. HPLC : Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 18.87 min, 22.01 min.

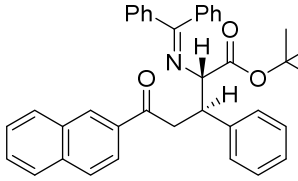
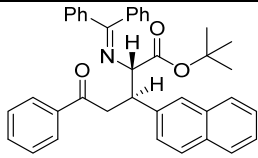
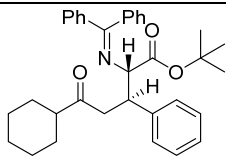
 <p>3.3, 90%, d.r. >99:1</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.3 (24.5 mg, 0.102 mmol) as starting material, compound 3.3 was obtained as colorless solid (49 mg, 96%, mp 144.2 °C).^{13,14,15}</p> <p>¹H NMR (400 MHz, Acetone) δ 7.98 (d, J = 8.6 Hz, 2H), 7.73 – 7.63 (m, 2H), 7.58 (tt, J = 7.2, 1.2 Hz, 1H), 7.51 – 7.35 (m, 8H), 7.23 (d, J = 1.2 Hz, 4H), 6.95 (dd, J = 6.6, 2.9 Hz, 2H), 4.21 (q, J = 3.4 Hz, 2H), 3.81 – 3.58 (m, 2H), 3.64 (dd, J = 17.1, 3.3 Hz, 1H), 1.29 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 198.7, 172.1, 170.3, 141.9, 140.5, 138.4, 137.3, 133.9, 132.8, 131.5, 131.5, 129.7, 129.7, 129.6, 129.5, 129.2, 129.02, 129.0, 128.6, 81.8, 71.8, 45.5, 41.1, 28.2. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 10.84 min, 14.07 min.</p>
 <p>3.4, 98%, d.r. 99:1</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.4 (29 mg, 0.102 mmol) as starting material, compound 3.4 was obtained as clear oil (58 mg, 98%).^{12,13}</p> <p>¹H NMR (400 MHz, Acetone) δ 8.02 – 7.94 (m, 2H), 7.71 – 7.64 (m, 2H), 7.59 (t, J = 7.4 Hz, 1H), 7.53 – 7.25 (m, 10H), 7.18 (d, J = 8.5 Hz, 2H), 6.94 (dd, J = 6.5, 3.0 Hz, 2H), 4.25 – 4.09 (m, 2H), 3.81 – 3.57 (m, 2H), 1.29 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 171.0, 169.2, 141.3, 139.4, 137.3, 136.2, 132.8, 130.9, 130.8, 130.4, 128.6, 128.6, 128.5, 128.4, 128.1, 127.9, 127.5, 119.8, 80.7, 70.7, 44.4, 39.9, 27.1. HPLC: Diacel IC (5μm), Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 70.82 min, 89.60 min.</p>
 <p>3.5, 79%, d.r. 98:2</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.5 (29 mg, 0.102 mmol) as starting material, compound 3.5 was obtained as clear oil (46 mg, 79%).¹⁵</p> <p>¹H NMR (500 MHz, acetone) δ 7.98 (d, J = 8.4 Hz, 1H), 7.68 (d, J = 8.5 Hz, 2H), 7.59 (t, J = 7.4 Hz, 1H), 7.52 – 7.42 (m, 6H), 7.41 – 7.34 (m, 4H), 7.18 (d, J = 8.5 Hz, 1H), 6.94 (dd, J = 6.4, 3.0 Hz, 2H), 4.23 – 4.15 (m, 2H), 3.80 – 3.71 (m, 1H), 3.68 – 3.59 (m, 1H), 1.29 (s, 9H). ¹³C NMR (126 MHz, acetone) δ 198.7, 172.1, 170.3, 142.4, 140.5, 138.3, 137.3, 133.9, 132.0, 131.9, 131.5, 129.7, 129.7, 129.6, 129.5, 129.2, 129.0, 128.6, 120.9, 81.8, 71.8, 45.5, 41.0. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 10.95 min, 14.41 min.</p>
 <p>3.6, 97%, d.r. 99:1</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.6 (22.6 mg, 0.102 mmol) as starting material, compound 3.6 was obtained as clear oil (51 mg, 97%).</p> <p>¹H NMR (500 MHz, acetone) δ 8.11 – 8.03 (m, 2H), 7.68 (d, J = 7.1 Hz, 1H), 7.47 – 7.41 (m, 4H), 7.38 (t, J = 7.4 Hz, 2H), 7.26 – 7.11 (m, 7H), 6.88 (dd, J = 5.0, 2.5 Hz, 2H), 4.27 – 4.19 (m, 1H), 3.79 – 3.69 (m, 1H), 3.68 – 3.61 (m, 1H), 1.28 (s, 9H). ¹³C NMR (126 MHz, acetone) δ 196.4, 170.8,</p>

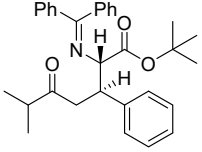
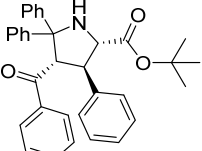
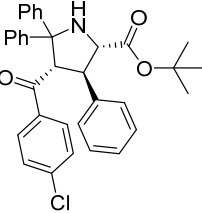
	<p>169.4, 165.4 (d, $J = 251.8$ Hz), 141.8, 139.5, 136.3, 134.0 (d, $J = 2.9$ Hz), 130.9 (d, $J = 9.4$ Hz), 130.4, 128.7, 128.61, 128.5, 128.3, 128.0, 127.97, 127.6, 126.5, 115.4 (d, $J = 22.0$ Hz), 80.5, 71.0, 45.0, 40.1, 27.1. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 12.58 min, 14.56 min.</p>
 <p>3.7, 97%, d.r. 98:2</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.7 (22.6 mg, 0.102 mmol) as starting material, compound 3.7 was obtained as clear oil (51 mg, 97%).^{13,14,15}</p> <p>¹H NMR (400 MHz, Acetone) δ 7.98 (d, $J = 8.5$ Hz, 2H), 7.73 – 7.65 (m, 2H), 7.58 (t, $J = 7.4$ Hz, 1H), 7.52 – 7.42 (m, 6H), 7.42 – 7.34 (m, 2H), 7.28 – 7.20 (m, 2H), 7.01 – 6.89 (m, 4H), 4.29 – 4.17 (m, 2H), 3.78 – 3.58 (m, 2H), 1.29 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 198.9, 172.1, 170.4, 140.6, 138.9, 138.5, 137.4, 133.9, 131.6, 131.6, 131.5, 129.7, 129.7, 129.6, 129.5, 129.2, 129.1, 128.7, 115.6 (d, $J = 22.0$ Hz), 81.7, 72.1, 45.4, 41.4, 28.2. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 11.21 min, 15.38 min.</p>
 <p>3.8, 79%, d.r. >99:1</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.8 (28 mg, 0.102 mmol) as starting material, compound 3.8 was obtained as colorless foam (45 mg, 79%, 96.2 °C).¹²</p> <p>¹H NMR (500 MHz, acetone) δ 7.86 (d, $J = 7.7$ Hz, 2H), 7.55 (d, $J = 7.5$ Hz, 2H), 7.45 (t, $J = 7.4$ Hz, 1H), 7.41 (d, $J = 8.2$ Hz, 2H), 7.38 – 7.27 (m, 9H), 7.25 (t, $J = 7.6$ Hz, 2H), 6.76 (d, $J = 6.3$ Hz, 2H), 4.20 – 4.13 (m, 1H), 4.11 (d, $J = 6.0$ Hz, 1H), 3.73 (dd, $J = 17.5, 9.9$ Hz, 1H), 3.59 (dd, $J = 17.5, 3.8$ Hz, 1H), 1.15 (s, 9H). ¹³C NMR (126 MHz, acetone) δ 197.5, 171.2, 169.1, 146.7, 139.4, 137.2, 136.1, 132.9, 130.5, 129.5, 128.7, 128.6, 128.6, 128.4, 128.1, 127.9, 127.5, 124.8 (q, $J = 3.8$ Hz), 80.9, 70.6, 44.7, 39.7, 27.1. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 9.98 min, 12.51 min.</p>
 <p>3.9, 70%, d.r. 83:17</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.9 (23.3 mg, 0.102 mmol) as starting material, compound 3.8 was obtained as colorless foam (37 mg, 70%).</p> <p>¹H NMR (400 MHz, Acetone) δ 7.98 (d, $J = 7.4$ Hz, 2H), 7.67 (d, $J = 7.2$ Hz, 2H), 7.59 (t, $J = 8.2$ Hz, 3H), 7.51 – 7.41 (m, 8H), 7.38 (t, $J = 7.4$ Hz, 2H), 6.92 (d, $J = 3.5$ Hz, 2H), 4.31 – 4.20 (m, 2H), 3.86 (dd, $J = 17.5, 9.4$ Hz, 1H), 3.70 (dd, $J = 17.5, 3.7$ Hz, 1H), 1.29 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 197.5, 169.0, 147.8, 139.3, 137.2, 136.1, 132.9, 131.7, 130.5, 129.9, 129.7, 129.3, 128.7, 128.6, 128.4, 128.1, 127.9, 127.5, 110.3, 80.9, 70.3, 44.9, 39.7, 27.1. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 22.04 min, 30.87 min. HRMS (m/z) calculated: C₃₅H₃₃N₂O₃ [M+ H]⁺: 529.2491, found: 529.2492.</p>

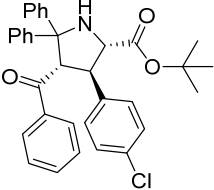
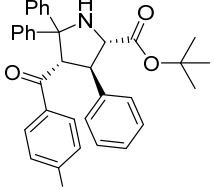
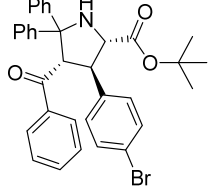
 <p>3.10, 71%, d.r. 98:2</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.10 (22.3 mg, 0.102 mmol) as starting material, compound 3.10 was obtained as clear oil (37.5 mg, 71%).¹³</p> <p>¹H NMR (500 MHz, acetone) δ 7.88 (d, J = 8.2 Hz, 2H), 7.68 (d, J = 8.6 Hz, 1H), 7.49 – 7.35 (m, 6H), 7.28 (d, J = 8.0 Hz, 2H), 7.21 – 7.09 (m, 5H), 6.92 – 6.84 (m, 2H), 4.25 – 4.17 (m, 2H), 3.75 – 3.67 (m, 1H), 3.61 – 3.53 (m, 1H), 2.37 (s, 3H), 1.28 (s, 9H). ¹³C NMR (126 MHz, acetone) δ 198.4, 171.8, 170.5, 144.5, 142.9, 140.6, 137.4, 136.0, 131.4, 130.2, 129.8, 129.7, 129.6, 129.4, 129.2, 129.1, 128.9, 128.6, 127.5, 81.5, 72.2, 46.1, 41.0, 28.2, 21.7. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 12.48 min, 17.20 min.</p>
 <p>3.11, 92%, d.r. 94:6</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.11 (22.3 mg, 0.102 mmol) as starting material, compound 3.11 was obtained as clear oil (48 mg, 92%).¹³</p> <p>¹H NMR (400 MHz, Acetone) δ 7.98 (d, J = 7.1 Hz, 2H), 7.68 (d, J = 8.6 Hz, 2H), 7.57 (t, J = 7.4 Hz, 1H), 7.51 – 7.35 (m, 8H), 7.06 (d, J = 8.1 Hz, 2H), 6.99 (d, J = 8.0 Hz, 2H), 6.92 – 6.84 (m, 2H), 4.26 – 4.11 (m, 2H), 3.78 – 3.65 (m, 1H), 3.60 (dd, J = 16.4, 3.2 Hz, 1H), 2.22 (s, 3H), 1.29 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 198.9, 171.7, 170.5, 140.6, 139.8, 138.5, 137.4, 136.9, 133.7, 131.4, 129.63, 129.61, 129.6, 129.5, 129.5, 129.4, 129.1, 129.9, 128.6, 81.5, 72.2, 45.6, 41.2, 28.2, 21.1. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 10.62 min, 14.11 min.</p>
 <p>3.12, 85%, d.r. >99:1</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.12 (24 mg, 0.102 mmol) as starting material, compound 3.12 was obtained as clear oil (45 mg, 85%).^{13,15}</p> <p>¹H NMR (400 MHz, Acetone) δ 7.99 – 7.92 (m, 2H), 7.72 – 7.63 (m, 2H), 7.48 – 7.33 (m, 6H), 7.20 – 7.08 (m, 5H), 7.00 – 6.94 (m, 2H), 6.91 – 6.83 (m, 2H), 4.26 – 4.14 (m, 2H), 3.85 (s, 3H), 3.72 – 3.61 (m, 1H), 3.57 – 3.48 (m, 1H), 1.27 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 196.2, 170.7, 169.4, 163.4, 141.9, 139.6, 136.4, 130.4, 130.3, 130.2, 128.7, 128.6, 128.5, 128.3, 128.0, 127.9, 127.6, 126.4, 113.6, 80.4, 71.1, 54.9, 45.1, 39.7, 27.1. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 20.67 min, 33.78 min.</p>
 <p>3.113, 88%, d.r. 98:2</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.13 (24 mg, 0.102 mmol) as starting material, compound 3.12 was obtained as clear oil (47 mg, 88%).^{13,15}</p>

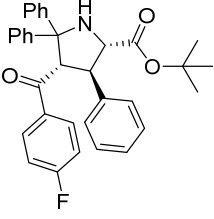
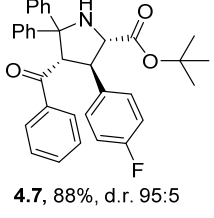
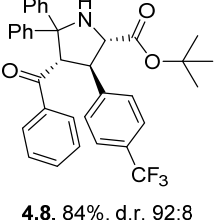
	<p>¹H NMR (400 MHz, Acetone) δ 7.97 (d, J = 7.4 Hz, 2H), 7.69 (d, J = 7.3 Hz, 2H), 7.57 (d, J = 8.0 Hz, 1H), 7.51 – 7.30 (m, 8H), 7.10 (d, J = 8.0 Hz, 2H), 6.99 – 6.86 (m, 2H), 6.74 (d, J = 7.9 Hz, 2H), 4.27 – 4.09 (m, 2H), 3.80 – 3.48 (m, 1H, c), 3.70 (s, 3H), 1.29 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 199.0, 171.7, 170.6, 159.6, 140.7, 138.6, 137.5, 134.7, 133.8, 131.4, 130.8, 130.7, 129.7, 129.6, 129.4, 129.2, 129.0, 128.71, 114.4, 81.5, 72.4, 55.6, 45.4, 41.5, 28.3. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 15.67 min, 25.88 min.</p>
 <p>3.14, 89%, d.r. >99:1</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.14 (24.5 mg, 0.102 mmol) as starting material, compound 3.14 was obtained as clear oil (48 mg, 89%).^{13,14,15}</p> <p>¹H NMR (500 MHz, acetone) δ 8.04 (d, J = 8.4 Hz, 1H), 7.70 – 7.64 (m, 2H), 7.63 – 7.58 (m, 1H), 7.51 (dd, J = 10.5, 4.8 Hz, 2H), 7.47 – 7.29 (m, 8H), 7.17 (td, J = 7.7, 1.7 Hz, 1H), 7.12 (td, J = 7.5, 1.2 Hz, 1H), 6.61 (d, J = 6.6 Hz, 2H), 4.77 (dt, J = 10.0, 4.1 Hz, 1H), 4.30 (d, J = 4.3 Hz, 1H), 4.10 – 4.02 (m, 1H), 3.77 (dd, J = 17.4, 4.0 Hz, 1H), 1.36 (d, J = 13.5 Hz, 9H). ¹³C NMR (126 MHz, acetone) δ 198.6, 172.1, 170.4, 140.4, 140.2, 138.3, 137.3, 135.3, 133.9, 131.4, 130.6, 130.5, 129.7, 129.6, 129.5, 129.3, 129.1, 129.0, 128.8, 128.3, 127.5, 81.9, 69.2, 41.6, 39.6, 28.3. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 11.95 min, 12.11 min.</p>
 <p>3.15, 91%, d.r. >99:1</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.15 (24.5 mg, 0.102 mmol) as starting material, compound 3.15 was obtained as clear oil (49 mg, 91%).¹⁴</p> <p>¹H NMR (500 MHz, acetone) δ 8.00 (d, J = 7.2 Hz, 2H), 7.68 (d, J = 7.1 Hz, 2H), 7.59 (t, J = 7.4 Hz, 1H), 7.52 – 7.42 (m, 6H), 7.39 (t, J = 7.4 Hz, 2H), 7.27 (s, 1H), 7.24 – 7.15 (m, 3H), 6.93 (dd, J = 6.4, 2.9 Hz, 2H), 4.25 – 4.17 (m, 2H), 3.84 – 3.76 (m, 1H), 3.71 – 3.62 (m, 1H), 1.29 (s, 9H). ¹³C NMR (126 MHz, acetone) δ 197.6, 171.1, 169.2, 144.4, 139.4, 137.2, 136.2, 133.3, 132.9, 130.4, 129.6, 128.9, 128.62, 128.58, 128.5, 128.4, 128.0, 127.9, 127.5, 127.1, 126.5, 80.73, 70.6, 44.7, 39.8, 27.1. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 10.96 min, 12.71 min.</p>
 <p>3.16, 88%, d.r. >99:1</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.16 (23.8 mg, 0.102 mmol) as starting material, compound 3.16 was obtained as clear oil (48 mg, 88%).</p> <p>¹H NMR (400 MHz, Acetone) δ 7.98 (d, J = 7.3 Hz, 2H), 7.66 (d, J = 7.2 Hz, 2H), 7.58 (t, J = 7.3 Hz, 1H), 7.52 – 7.34 (m, 8H), 7.16 – 7.07 (m, 2H), 6.87 – 6.76 (m, 3H), 6.73 (t, J = 7.4 Hz, 1H), 4.57 – 4.49 (m, 1H), 4.45 (d, J = 6.0 Hz, 1H), 3.87 – 3.75 (m, 1H), 3.62 – 3.49 (m, 4H), 1.28 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 198.2, 170.5, 169.9, 157.6, 139.6, 137.6, 136.6,</p>

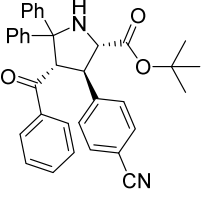
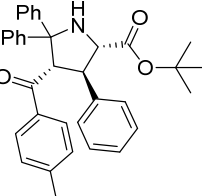
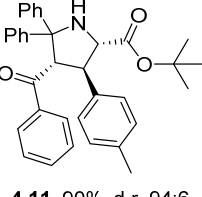
	132.6, 130.2, 129.5, 129.3, 128.6, 128.4, 128.3, 128.2, 128.0, 127.9, 127.6, 127.5, 119.8, 110.9, 80.1, 68.3, 54.6, 39.7, 38.5, 27.1. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 17.31 min, 20.64 min. HRMS (m/z) calculated: C ₃₅ H ₃₆ NO ₄ [M+H] ⁺ : 534.2644, found: 534.2636.
 <p>3.17, 90%, d.r. 98:2</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.17 (23.8 mg, 0.102 mmol) as starting material, compound 3.17 was obtained as clear oil (48 mg, 90%).¹⁴</p> <p>¹H NMR (500 MHz, acetone) δ 8.00 (d, J = 7.2 Hz, 1H), 7.70 (d, J = 7.1 Hz, 1H), 7.58 (t, J = 7.4 Hz, 1H), 7.52 – 7.35 (m, 4H), 7.09 (t, J = 8.1 Hz, 1H), 6.91 – 6.81 (m, 1H), 6.79 – 6.73 (m, 1H), 6.74 – 6.67 (m, 1H), 4.26 – 4.16 (m, 1H), 3.84 – 3.73 (m, 1H), 3.69 – 3.64 (m, 1H), 3.65 (s, 3H), 1.31 (s, 9H). ¹³C NMR (126 MHz, acetone) δ 198.9, 171.7, 170.5, 160.6, 144.5, 140.6, 138.5, 137.4, 133.8, 131.4, 129.9, 129.7, 129.6, 129.6, 129.4, 129.1, 129.0, 128.6, 121.9, 115.5, 113.0, 81.6, 72.0, 55.5, 45.9, 40.9, 28.2. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 13.15 min, 16.18 min.</p>
 <p>3.18, 90%, d.r. 96:4</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.18 (21.4 mg, 0.102 mmol) as starting material, compound 3.18 was obtained as clear oil (46 mg, 90%).</p> <p>¹H NMR (400 MHz, Acetone) δ 8.30 (s, 1H), 8.01 – 7.96 (m, 1H), 7.96 – 7.91 (m, 1H), 7.71 – 7.66 (m, 2H), 7.52 – 7.31 (m, 8H), 7.25 – 7.10 (m, 5H), 6.88 (dd, J = 7.0, 2.5 Hz, 2H), 4.31 – 4.18 (m, 2H), 3.86 – 3.78 (m, 1H), 3.72 – 3.63 (m, 1H), 1.29 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 192.6, 170.9, 169.3, 144.1, 142.2, 141.6, 139.5, 136.3, 130.4, 129.7, 128.7, 128.6, 128.5, 128.4, 128.0, 127.6, 127.4, 126.6, 126.1, 124.9, 122.8, 80.6, 70.9, 45.2, 40.6, 27.1. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 18.81 min, 22.75 min. HRMS (m/z) calculated: C₃₆H₃₄NO₃S [M+H]⁺: 560.2259, found: 560.2275</p>
 <p>3.19, 91%, d.r. >99:1</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.19 (19.82 mg, 0.102 mmol) as starting material, compound 3.19 was obtained as colorless solid (45 mg, 91%, mp 89.9 °C).^{12,13,14}</p> <p>¹H NMR (500 MHz, acetone) δ 7.77 – 7.72 (m, 1H), 7.70 – 7.64 (m, 2H), 7.48 – 7.41 (m, 4H), 7.38 (t, J = 7.4 Hz, 2H), 7.27 (d, J = 3.5 Hz, 1H), 7.22 – 7.10 (m, 5H), 6.91 – 6.82 (m, 2H), 6.58 (dd, J = 3.5, 1.7 Hz, 1H), 4.25 – 4.16 (m, 2H), 3.64 – 3.55 (m, 1H), 3.40 (dd, J = 16.3, 3.7 Hz, 1H), 1.29 (s, 9H). ¹³C NMR (126 MHz, acetone) δ 186.6, 170.8, 169.3, 153.0, 146.6, 141.6, 139.5, 136.3, 130.4, 128.7, 128.6, 128.5, 128.34, 128.0, 127.9, 127.6, 126.5, 116.9, 112.1, 80.5, 71.0, 44.8, 39.9, 27.1. HPLC: Kromasil</p>

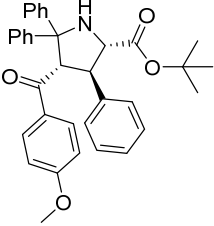
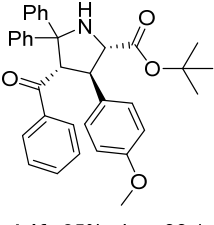
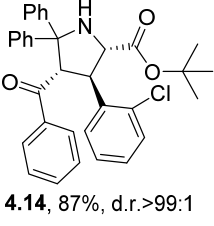
	OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 19.63 min, 26.06 min.
 <p>3.20, 87%, d.r. >99:1</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.20 (23.8 mg, 0.102 mmol) as starting material, compound 3.20 was obtained as clear oil (48 mg, 87%).</p> <p>$^1\text{H NMR}$ (400 MHz, Acetone) δ 8.66 (s, 1H), 8.09 (d, $J = 8.0$ Hz, 1H), 8.04 – 7.88 (m, 3H), 7.73 – 7.53 (m, 5H), 7.48 – 7.34 (m, 6H), 7.27 – 7.11 (m, 5H), 6.97 – 6.83 (m, 2H), 4.34 – 4.22 (m, 2H), 3.88 (dd, $J = 16.8, 9.0$ Hz, 1H), 3.77 (dd, $J = 16.8, 4.1$ Hz, 1H), 1.29 (s, 11H). $^{13}\text{C NMR}$ (101 MHz, Acetone) δ 198.9, 171.9, 170.5, 143.0, 140.7, 137.5, 136.6, 135.9, 133.8, 131.4, 130.8, 130.8, 130.7, 129.9, 129.7, 129.6, 129.4, 129.3, 129.1, 129.1, 128.8, 128.7, 127.8, 127.6, 124.9, 81.6, 72.2, 46.3, 41.3, 28.2. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 16.09 min, 17.96 min.</p>
 <p>3.21, 83%, d.r. >99:1</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.21 (23.8 mg, 0.102 mmol) as starting material, compound 3.21 was obtained as clear oil (46 mg, 83%).</p> <p>$^1\text{H NMR}$ (400 MHz, Acetone) δ 8.17 (d, $J = 8.1$ Hz, 1H), 8.04 (dd, $J = 8.3, 1.3$ Hz, 2H), 7.88 (d, $J = 7.6$ Hz, 1H), 7.74 (d, $J = 8.1$ Hz, 1H), 7.69 – 7.63 (m, 2H), 7.59 (t, $J = 8.0$ Hz, 1H), 7.52 – 7.33 (m, 9H), 7.33 – 7.25 (m, 2H), 7.21 (t, $J = 7.5$ Hz, 1H), 7.08 (t, $J = 7.6$ Hz, 2H), 6.41 (d, $J = 6.3$ Hz, 2H), 5.30 – 5.14 (m, 1H), 4.34 (d, $J = 4.5$ Hz, 1H), 4.14 (dd, $J = 17.4, 9.4$ Hz, 1H), 3.86 (dd, $J = 17.4, 4.3$ Hz, 1H), 1.30 (s, 9H). $^{13}\text{C NMR}$ (101 MHz, Acetone) δ 198.9, 171.8, 170.8, 140.5, 138.8, 138.5, 137.0, 135.2, 133.8, 133.0, 131.3, 129.7, 129.6, 129.6, 129.2, 129.0, 129.0, 128.9, 128.0, 128.0, 126.9, 126.4, 125.9, 124.2, 81.7, 70.6, 40.4, 28.2. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 13.28 min, 14.93 min.</p>
 <p>3.22, 96%, d.r. 98:2</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.22 (21.5 mg, 0.102 mmol) as starting material, compound 3.22 was obtained as clear oil (49 mg, 96%).</p> <p>$^1\text{H NMR}$ (500 MHz, acetone) δ 7.52 (d, $J = 8.6$ Hz, 1H), 7.33 – 7.21 (m, 1H), 7.09 – 6.97 (m, 5H), 6.70 (d, $J = 7.5$ Hz, 1H), 3.97 (d, $J = 6.0$ Hz, 1H), 3.90 – 3.85 (m, 1H), 3.06 (dd, $J = 17.1, 10.1$ Hz, 1H), 2.91 (dd, $J = 17.1, 3.9$ Hz, 1H), 2.21 (ddd, $J = 11.1, 7.3, 3.3$ Hz, 1H), 1.63 – 1.39 (m, 6H), 1.15 (s, 9H), 1.12 – 1.03 (m, 2H), 1.02 – 0.87 (m, 2H). $^{13}\text{C NMR}$ (126 MHz, acetone) δ 210.5, 170.5, 169.4, 142.0, 139.5, 136.3, 130.3, 128.7, 128.5, 128.5, 128.3, 128.1, 127.9, 127.5, 126.4, 80.4, 70.9, 50.3, 44.3, 42.1, 28.1, 28.1, 27.2, 25.7, 25.4, 25.3. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 9.26 min, 13.36 min.</p>

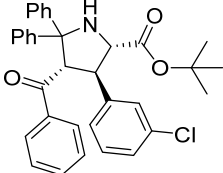
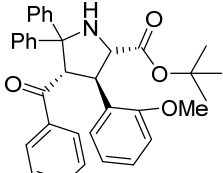
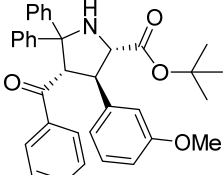
	<p>HRMS (m/z) calculated: C₃₄H₄₀NO₃ [M+H]⁺: 510.3008, found: 510.3003</p>
 <p>3.23, 70%, d.r. 98:2</p>	<p>Following Procedure C using α,β-unsaturated ketone 2.23 (23.8 mg, 0.102 mmol) as starting material, compound 3.22 was obtained as clear oil (34 mg, 70%).</p> <p>¹H NMR (500 MHz, acetone) δ 7.66 (d, J = 7.1 Hz, 2H), 7.52 – 7.32 (m, 6H), 7.26 – 7.08 (m, 5H), 6.85 (d, J = 7.4 Hz, 2H), 4.11 (d, J = 6.1 Hz, 1H), 4.01 (ddd, J = 10.1, 6.0, 4.0 Hz, 1H), 3.20 (dd, J = 17.1, 10.1 Hz, 1H), 3.06 (dd, J = 17.1, 3.9 Hz, 1H), 2.59 (dq, J = 13.8, 6.9 Hz, 1H), 1.28 (s, 9H), 0.95 (d, J = 6.9 Hz, 3H), 0.87 (d, J = 7.0 Hz, 3H). ¹³C NMR (126 MHz, acetone) δ 211.3, 170.6, 169.3, 141.9, 139.5, 136.3, 130.4, 128.7, 128.5, 128.5, 128.3, 128.1, 127.9, 127.5, 126.4, 80.4, 70.9, 44.4, 41.9, 40.5, 27.1, 17.37, 17.35. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 8.99 min, 12.06 min. HRMS (m/z) calculated: C₃₁H₃₆NO₃ [M+H]⁺: 470.2695, found: 470.2699</p>
 <p>4.1, 95%, d.r. 96:4</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.1 (21 mg, 0.102 mmol) as starting material, compound 3.22 was obtained as clear oil (49 mg, 95%).</p> <p>¹H NMR (400 MHz, Acetone) δ 7.95 (d, J = 7.5 Hz, 2H), 7.73 (d, J = 7.3 Hz, 2H), 7.57 (d, J = 7.5 Hz, 1H), 7.48 – 7.31 (m, 6H), 7.27 (t, J = 7.3 Hz, 1H), 7.21 – 7.12 (m, 3H), 7.10 – 6.95 (m, 3H), 6.86 (dd, J = 6.7, 2.8 Hz, 2H), 5.42 (d, J = 4.8 Hz, 1H), 4.01 (d, J = 12.4 Hz, 1H), 3.93 – 3.82 (m, 1H), 3.71 (dd, J = 8.8, 4.8 Hz, 1H), 1.35 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 203.4, 172.3, 147.6, 146.6, 142.3, 139.7, 134.0, 130.8, 129.8, 129.75, 129.5, 129.2, 129.1, 128.9, 128.0, 127.94, 127.91, 127.5, 127.2, 81.9, 78.4, 68.9, 64.5, 60.3, 28.3. HPLC: Kromasil OD-H, Hexane:2-Propanol (96:4), 0.5 mL/min, 254 nm; retention time: 10.21 min, 13.86 min. HRMS (m/z) calculated: C₃₄H₃₃NO₃Na [M+Na]⁺: 526.2358, found: 526.2339.</p>
 <p>4.2, 96%, d.r. 96:4</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.2 (24.5 mg, 0.102 mmol) as starting material, compound 4.2 was obtained as clear oil (52 mg, 96%).</p> <p>¹H NMR (400 MHz, Acetone) δ 7.94 (d, J = 7.4 Hz, 2H), 7.72 (d, J = 8.6 Hz, 2H), 7.45 – 7.31 (m, 7H), 7.26 (t, J = 7.3 Hz, 1H), 7.19 – 7.13 (m, 3H), 7.10 – 6.95 (m, 3H), 6.86 (dd, J = 6.4, 3.1 Hz, 2H), 5.38 (d, J = 4.7 Hz, 1H), 4.10 – 3.94 (m, 1H), 3.86 (dd, J = 13.3, 6.3 Hz, 1H), 3.73 (dd, J = 8.7, 4.7 Hz, 1H), 1.34 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 201.2, 171.1, 146.3, 145.2, 141.1, 138.8, 137.1, 129.8, 128.8, 128.7, 128.4, 128.1, 127.9, 127.0, 126.9, 126.9, 126.5, 126.2, 80.8, 77.3, 67.7, 63.4, 58.9, 27.2. HPLC: Diacel IC (5μm), Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 17.86 min, 20.46 min. HRMS (m/z) calculated: C₃₄H₃₂ClNO₃Na [M+Na]⁺: 560.1968, found: 560.1959.</p>

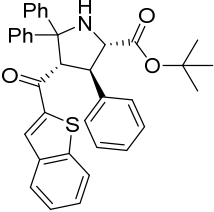
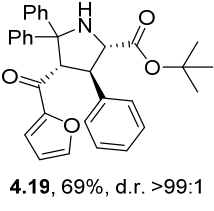
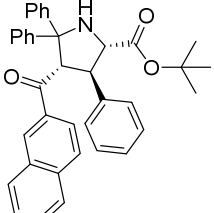
 <p>4.3, 92%, d.r. 90:10</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.3 (24.5 mg, 0.102 mmol) as starting material, compound 4.3 was obtained as clear oil (50 mg, 92%).</p> <p>¹H NMR (400 MHz, Acetone) δ 7.93 (d, J = 7.5 Hz, 2H), 7.74 (d, J = 7.4 Hz, 2H), 7.57 (t, J = 7.4 Hz, 1H), 7.45 – 7.31 (m, 6H), 7.26 (t, J = 7.3 Hz, 1H), 7.20 – 7.13 (m, 2H), 7.09 – 6.95 (m, 3H), 6.85 (d, J = 8.4 Hz, 2H), 5.39 (d, J = 4.6 Hz, 1H), 4.01 (d, J = 11.8 Hz, 1H), 3.93 – 3.79 (m, 1H), 3.72 (dd, J = 8.5, 4.6 Hz, 1H), 1.36 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 203.1, 172.0, 147.4, 146.3, 141.4, 139.6, 134.1, 133.2, 131.0, 130.8, 129.9, 129.8, 129.4, 129.2, 128.9, 128.1, 127.9, 127.5, 127.2, 82.1, 78.4, 68.7, 64.2, 59.0, 28.3. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 10.37 min, 15.74 min. HRMS (m/z) calculated: C₃₄H₃₂ClNO₃Na [M+Na]⁺: 560.1968, found: 560.1956.</p>
 <p>4.4, 76%, d.r. 90:10</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.4 (29 mg, 0.102 mmol) as starting material, compound 4.4 was obtained as clear oil (44 mg, 76%).</p> <p>¹H NMR (400 MHz, Acetone) δ 7.93 (d, J = 8.2 Hz, 2H), 7.74 (d, J = 8.3 Hz, 2H), 7.57 (t, J = 7.4 Hz, 1H), 7.45 – 7.30 (m, 8H), 7.26 (t, J = 7.3 Hz, 1H), 7.08 – 6.96 (m, 3H), 6.79 (d, J = 8.5 Hz, 2H), 5.39 (d, J = 4.6 Hz, 1H), 4.02 (d, J = 11.9 Hz, 1H), 3.86 (dd, J = 11.9, 8.6 Hz, 1H), 3.71 (dd, J = 8.6, 4.5 Hz, 1H), 1.36 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 202.0, 170.9, 146.3, 145.2, 140.8, 138.5, 133.0, 131.3, 130.2, 128.8, 128.7, 128.1, 127.8, 127.0, 126.9, 126.4, 126.2, 120.2, 81.0, 77.3, 67.6, 63.0, 57.9, 27.2. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 10.62 min, 16.37 min. HRMS (m/z) calculated: C₃₄H₃₂BrNO₃Na [M+Na]⁺: 604.1463, found: 604.1452.</p>
 <p>4.5, 86%, d.r. 90:10</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.5 (29 mg, 0.102 mmol) as starting material, compound 4.5 was obtained as clear oil (50 mg, 86%).</p> <p>¹H NMR (400 MHz, Acetone) δ 7.95 (d, J = 8.1 Hz, 2H), 7.75 (d, J = 7.7 Hz, 2H), 7.57 (d, J = 7.6 Hz, 1H), 7.47 – 7.23 (m, 9H), 7.10 – 6.94 (m, 3H), 6.81 (d, J = 8.3 Hz, 2H), 5.40 (d, J = 4.5 Hz, 1H), 4.03 (d, J = 10.0 Hz, 1H), 3.87 (d, J = 7.0 Hz, 1H), 3.73 (dd, J = 8.4, 4.4 Hz, 1H), 1.37 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 202.0, 170.9, 146.3, 145.2, 140.8, 138.5, 133.0, 131.3, 130.2, 128.8, 128.7, 128.1, 127.8, 127.0, 126.9, 126.4, 126.2, 120.2, 81.0, 77.3, 67.6, 63.0, 58.0, 27.3. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 10.68 min, 16.54 min. HRMS (m/z) calculated: C₃₄H₃₂BrNO₃Na [M+ Na]⁺: 604.1463, found: 604.1456.</p>

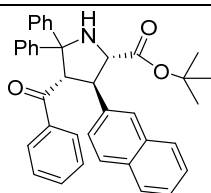
 <p>4.6, 96%, d.r. 96:4</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.6 (22.6 mg, 0.102 mmol) as starting material, compound 4.6 was obtained as colorless solid (50 mg, 96%, mp 138.2 °C).</p> <p>¹H NMR (400 MHz, Acetone) δ 7.96 (d, J = 8.2 Hz, 2H), 7.86 – 7.78 (m, 2H), 7.42 – 7.30 (m, 4H), 7.27 (t, J = 7.3 Hz, 1H), 7.21 – 7.10 (m, 5H), 7.09 – 6.96 (m, 3H), 6.90 – 6.79 (m, 2H), 5.40 (d, J = 4.7 Hz, 1H), 4.00 (d, J = 11.6 Hz, 1H), 3.87 (t, J = 9.3 Hz, 1H), 3.78 – 3.66 (m, 1H), 1.35 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 200.8, 171.2, 165.6 (d, J = 252.6 Hz), 146.4, 145.3, 141.2, 135.2, 131.0 (d, J = 9.4 Hz), 128.7, 128.4, 128.1, 127.8, 127.0, 126.9, 126.8, 126.4, 126.1, 115.54 (d, J = 22.1 Hz), 80.8, 77.3, 67.7, 63.3, 58.9, 27.2. HPLC: Diacel IC (5μm), Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 17.70 min, 20.61 min. HRMS (m/z) calculated: C₃₄H₃₂FNO₃Na [M+Na]⁺: 544.2264, found: 544.2260.</p>
 <p>4.7, 88%, d.r. 95:5</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.7 (22.6 mg, 0.102 mmol) as starting material, compound 4.7 was obtained as colorless solid (46 mg, 88%, mp 126.2 °C).</p> <p>¹H NMR (400 MHz, Acetone) δ 7.95 (d, J = 7.3 Hz, 2H), 7.74 (d, J = 7.2 Hz, 2H), 7.58 (t, J = 7.4 Hz, 1H), 7.39 (dddd, J = 8.2, 6.1, 5.1, 1.4 Hz, 6H), 7.27 (t, J = 7.3 Hz, 1H), 7.14 – 6.97 (m, 3H), 6.97 – 6.82 (m, 4H), 5.40 (d, J = 4.7 Hz, 1H), 4.09 – 3.94 (m, 1H), 3.92 – 3.82 (m, 1H), 3.74 (dd, J = 8.7, 4.6 Hz, 1H), 1.37 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 203.2, 172.1, 147.5, 146.5, 139.7, 138.5, 134.1, 131.1, 131.0, 129.9, 129.8, 129.5, 129.1, 128.9, 128.1, 127.9, 127.5, 127.2, 116.06 (d, J = 21.3 Hz), 101.1, 82.0, 78.3, 68.9, 64.4, 59.2, 28.3. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 10.59 min, 15.63 min. HRMS (m/z) calculated: C₃₄H₃₂FNO₃Na [M+Na]⁺: 544.2264, found: 544.2252.</p>
 <p>4.8, 84%, d.r. 92:8</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.8 (28 mg, 0.102 mmol) as starting material, compound 4.8 was obtained as clear oil (48 mg, 84%).</p> <p>¹H NMR (400 MHz, Acetone) δ 7.96 (dd, J = 8.4, 1.0 Hz, 2H), 7.79 – 7.73 (m, 2H), 7.57 (dd, J = 11.7, 4.4 Hz, 1H), 7.51 (d, J = 8.1 Hz, 2H), 7.47 – 7.32 (m, 6H), 7.29 (dt, J = 9.0, 4.2 Hz, 1H), 7.13 – 6.97 (m, 5H), 5.46 (d, J = 4.5 Hz, 1H), 4.05 (d, J = 11.4 Hz, 1H), 3.94 (dt, J = 11.5, 5.7 Hz, 1H), 3.86 (dd, J = 8.7, 4.4 Hz, 1H), 1.37 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 201.9, 170.8, 146.1, 146.0, 145.0, 138.5, 133.0, 128.9, 128.8, 128.7, 128.4, 128.1, 127.83, 127.1, 126.9, 126.5, 126.2, 125.2 (q, J = 3.9 Hz), 81.1, 77.3, 67.4, 62.8, 57.9, 27.2. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 9.71 min, 16.29 min. HRMS (m/z) calculated: C₃₅H₃₂F₃NO₃Na [M+Na]⁺: 594.2232, found: 594.2218.</p>

 <p>4.9, 70%, d.r. 75:25</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.9 (23.3 mg, 0.102 mmol) as starting material, compound 4.9 was obtained as clear oil (38 mg, 70%).</p> <p>¹H NMR (400 MHz, Acetone) δ 7.93 (d, J = 7.5 Hz, 2H), 7.74 (d, J = 7.7 Hz, 2H), 7.68 (d, J = 8.4 Hz, 1H), 7.57 (d, J = 8.3 Hz, 3H), 7.39 (ddd, J = 27.1, 13.1, 8.5 Hz, 6H), 7.27 (t, J = 7.3 Hz, 1H), 7.09 (d, J = 8.3 Hz, 2H), 7.03 (d, J = 7.7 Hz, 2H), 5.44 (d, J = 4.5 Hz, 1H), 4.04 (dd, J = 9.3, 4.8 Hz, 1H), 3.93 (dd, J = 13.3, 6.4 Hz, 1H), 3.89 – 3.81 (m, 1H), 1.36 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 201.7, 170.7, 146.9, 144.8, 133.1, 132.1, 131.7, 131.5, 130.7, 129.9, 129.3, 128.8, 128.7, 128.6, 128.1, 127.8, 126.9, 126.5, 126.3, 110.6, 81.2, 77.4, 67.3, 62.7, 57.9, 27.2. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 18.32 min, 29.84 min. HRMS (m/z) calculated: C₃₅H₃₂N₂O₃Na [M+Na]⁺: 551.2311, found: 551.2295.</p>
 <p>4.10, 83%, d.r. 96:4</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.10 (22.3 mg, 0.102 mmol) as starting material, compound 4.10 was obtained as colorless solid (44 mg, 83%, mp 150.4 °C).</p> <p>¹H NMR (500 MHz, acetone) δ 7.96 (d, J = 7.5 Hz, 2H), 7.67 (d, J = 8.2 Hz, 2H), 7.42 – 7.34 (m, 4H), 7.29 – 7.21 (m, 3H), 7.20 – 7.11 (m, 3H), 7.06 (t, J = 7.5 Hz, 2H), 6.99 (t, J = 7.3 Hz, 1H), 6.87 – 6.78 (m, 2H), 5.40 (d, J = 4.6 Hz, 1H), 4.10 – 3.98 (m, 1H), 3.88 (t, J = 8.4 Hz, 1H), 3.65 (dd, J = 8.7, 4.6 Hz, 1H), 2.37 (s, 3H), 1.35 (s, 9H). ¹³C NMR (126 MHz, acetone) δ 202.9, 172.3, 147.7, 146.8, 144.9, 142.4, 137.2, 130.4, 129.8, 129.4, 129.3, 129.2, 128.9, 127.9, 127.8, 127.4, 127.1, 81.8, 78.4, 68.9, 64.2, 60.6, 28.3, 21.7. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 10.56 min, 11.42 min. HRMS (m/z) calculated: C₃₅H₃₆NO₃ [M+H]⁺: 518.2695, found: 518.2692</p>
 <p>4.11, 90%, d.r. 94:6</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.11 (22.3 mg, 0.102 mmol) as starting material, compound 4.11 was obtained as clear oil (47 mg, 90%).</p> <p>¹H NMR (400 MHz, Acetone) δ 7.94 (d, J = 7.4 Hz, 2H), 7.74 (d, J = 7.2 Hz, 2H), 7.58 (t, J = 7.4 Hz, 1H), 7.46 – 7.33 (m, 6H), 7.26 (t, J = 7.3 Hz, 1H), 7.06 (t, J = 7.3 Hz, 2H), 6.99 (dd, J = 14.0, 7.5 Hz, 3H), 6.72 (d, J = 7.9 Hz, 2H), 5.39 (d, J = 4.6 Hz, 1H), 4.11 – 3.94 (m, 1H), 3.91 – 3.78 (m, 1H), 3.67 (dd, J = 8.6, 4.5 Hz, 1H), 2.24 (s, 3H), 1.36 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 203.4, 172.2, 147.5, 146.6, 139.6, 139.2, 137.2, 133.9, 129.9, 129.65, 129.6, 128.95, 128.9, 128.8, 127.8, 127.8, 127.3, 127.0, 81.7, 78.2, 68.8, 64.4, 59.8, 28.2, 21.0. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 9.77 min, 13.64 min. HRMS (m/z) calculated: C₃₅H₃₅NO₃Na [M+Na]⁺: 540.2515, found: 540.2507.</p>

 <p>4.12, 92%, d.r. 95:5</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.12 (24 mg, 0.102 mmol) as starting material, compound 4.12 was obtained as colorless solid (50 mg, 92%, 147.9 °C).</p> <p>¹H NMR (400 MHz, Acetone) δ 7.96 (d, J = 7.4 Hz, 2H), 7.77 (d, J = 8.9 Hz, 2H), 7.42 – 7.30 (m, 4H), 7.26 (t, J = 7.3 Hz, 1H), 7.19 – 7.11 (m, 3H), 7.09 – 7.02 (m, 2H), 7.01 – 6.91 (m, 3H), 6.85 – 6.76 (m, 2H), 5.36 (d, J = 4.6 Hz, 1H), 4.02 (d, J = 12.3 Hz, 1H), 3.91 – 3.78 (m, 1H, covered), 3.87 (s, 3H), 3.63 (dd, J = 8.7, 4.6 Hz, 1H), 1.34 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 201.6, 172.3, 164.8, 147.8, 146.9, 142.5, 132.5, 131.5, 129.8, 129.4, 129.1, 128.83, 127.87, 127.85, 127.8, 127.3, 127.0, 114.9, 81.8, 78.3, 68.9, 63.9, 60.6, 56.1, 28.3. HPLC: Diacel IC (5μm), Hexane:2-Propanol (95:5), 1 mL/min, 254 nm; retention time: 22.09 min, 27.45 min. HRMS (m/z) calculated: C₃₅H₃₅NO₄Na [M+Na]⁺: 556.2464, found: 556.2462.</p>
 <p>4.13, 85%, d.r. >99:1</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.13 (24 mg, 0.102 mmol) as starting material, compound 4.11 was obtained as clear oil (48 mg, 85%).</p> <p>¹H NMR (400 MHz, Acetone) δ 7.95 (d, J = 7.4 Hz, 2H), 7.74 (d, J = 7.1 Hz, 2H), 7.62 – 7.55 (m, 1H), 7.47 – 7.31 (m, 6H), 7.26 (t, J = 7.3 Hz, 1H), 7.12 – 6.95 (m, 3H), 6.73 (q, J = 9.0 Hz, 4H), 5.37 (d, J = 4.6 Hz, 1H), 3.82 (d, J = 8.7 Hz, 1H), 3.73 (s, 3H), 3.64 (dd, J = 8.8, 4.6 Hz, 1H), 1.33 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 202.4, 171.3, 158.7, 146.6, 145.7, 138.6, 133.2, 132.9, 129.1, 128.7, 128.7, 128.0, 127.8, 126.9, 126.8, 126.3, 126.0, 113.7, 80.7, 77.2, 67.9, 63.6, 58.7, 54.5, 27.2. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 12.62 min, 20.11 min. HRMS (m/z) calculated: C₃₅H₃₅NO₄Na [M+ Na]⁺: 556.2464, found: 556.2454.</p>
 <p>4.14, 87%, d.r. >99:1</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.14 (24.5 mg, 0.102 mmol) as starting material, compound 4.14 was obtained as colorless solid (47 mg, 87%, mp 84.2 °C).</p> <p>¹H NMR (400 MHz, Acetone) δ 7.93 (d, J = 7.4 Hz, 2H), 7.77 (d, J = 7.3 Hz, 2H), 7.56 (t, J = 7.4 Hz, 1H), 7.45 – 7.24 (m, 8H), 7.19 (td, J = 7.7, 1.5 Hz, 1H), 7.08 – 6.92 (m, 4H), 6.49 (d, J = 8.0 Hz, 1H), 5.33 (d, J = 4.0 Hz, 1H), 4.37 (dd, J = 8.6, 4.0 Hz, 1H), 4.10 (d, J = 12.3 Hz, 1H), 3.97 (dd, J = 12.2, 8.6 Hz, 1H), 1.34 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 203.7, 171.8, 147.2, 145.4, 140.0, 139.7, 135.1, 133.9, 130.1, 129.8, 129.6, 129.5, 129.2, 129.1, 128.8, 128.2, 128.1, 128.0, 127.3, 127.2, 81.9, 78.6, 67.4, 63.3, 55.0, 28.1. HPLC: Diacel IC (5μm), Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 9.93 min, 10.99 min. HRMS (m/z) calculated: C₃₄H₃₃ClNO₃ [M+H]⁺: 538.2149, found: 538.2143.</p>

 <p>4.15, 87%, d.r. 93:7</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.15 (24.5 mg, 0.102 mmol) as starting material, compound 4.15 was obtained as colorless solid (47 mg, 87%, mp 112.9 °C).</p> <p>¹H NMR (500 MHz, acetone) δ 7.98 (d, J = 7.8 Hz, 2H), 7.78 (d, J = 7.5 Hz, 2H), 7.59 (t, J = 7.4 Hz, 1H), 7.48 – 7.34 (m, 6H), 7.28 (t, J = 7.3 Hz, 1H), 7.24 – 7.16 (m, 2H), 7.06 (t, J = 7.5 Hz, 2H), 7.00 (t, J = 7.3 Hz, 1H), 6.87 (d, J = 6.8 Hz, 1H), 6.78 (d, J = 0.8 Hz, 1H), 5.43 (d, J = 4.3 Hz, 1H), 4.09 – 3.99 (m, 1H), 3.90 – 3.83 (m, 1H), 3.71 (dd, J = 8.5, 4.2 Hz, 1H), 1.38 (s, 9H). ¹³C NMR (126 MHz, acetone) δ 202.8, 171.7, 147.0, 146.1, 144.7, 139.2, 134.6, 133.9, 130.7, 129.6, 129.5, 128.9, 128.9, 128.7, 127.9, 127.7, 127.7, 127.6, 127.2, 126.9, 81.8, 78.3, 68.7, 63.7, 59.0, 28.0. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 9.98 min, 13.17 min. HRMS (m/z) calculated: C₃₄H₃₃ClNO₃ [M+H]⁺: 538.2149, found: 538.2137.</p>
 <p>4.16, 87%, d.r. 98:2</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.16 (23.8 mg, 0.102 mmol) compound 4.16 was obtained as clear oil (46 mg, 87%).</p> <p>¹H NMR (500 MHz, acetone) δ 7.78 (dd, J = 19.2, 7.3 Hz, 4H), 7.56 (t, J = 7.4 Hz, 1H), 7.43 (t, J = 7.7 Hz, 2H), 7.35 – 7.24 (m, 4H), 7.19 (ddd, J = 17.2, 11.4, 4.5 Hz, 2H), 7.03 – 6.92 (m, 3H), 6.89 (d, J = 8.1 Hz, 1H), 6.73 – 6.67 (m, 1H), 6.63 (dd, J = 7.6, 1.3 Hz, 1H), 5.30 (d, J = 4.5 Hz, 1H), 4.14 (dd, J = 8.6, 4.5 Hz, 1H), 4.01 (d, J = 8.4 Hz, 1H), 3.38 (s, 3H), 1.33 (s, 9H). ¹³C NMR (126 MHz, acetone) δ 202.9, 171.4, 157.1, 146.5, 144.9, 138.9, 132.5, 129.0, 128.39, 128.38, 128.0, 127.7, 127.6, 127.3, 126.8, 126.7, 126.4, 125.9, 120.0, 110.1, 80.5, 76.7, 63.8, 61.5, 54.3, 51.7, 27.2. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 12.00 min, 17.33 min. HRMS (m/z) calculated: C₃₅H₃₆NO₄ [M+H]⁺: 534.2644, found: 534.2644.</p>
 <p>4.17, 84%, d.r. 85:15</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.17 (23.8 mg, 0.102 mmol) as starting material, compound 4.17 was obtained as colorless solid (44.5 mg, 84%, mp 107.8 °C).</p> <p>¹H NMR (400 MHz, Acetone) δ 7.99 (d, J = 7.4 Hz, 2H), 7.75 (d, J = 8.5 Hz, 1H), 7.59 (t, J = 7.4 Hz, 1H), 7.51 – 7.31 (m, 6H), 7.25 (t, J = 7.3 Hz, 1H), 7.07 (dd, J = 15.4, 7.9 Hz, 3H), 6.99 (t, J = 7.2 Hz, 1H), 6.73 (dd, J = 8.1, 2.2 Hz, 1H), 6.49 (d, J = 7.6 Hz, 1H), 6.43 – 6.33 (m, 1H), 5.44 (d, J = 4.5 Hz, 1H), 4.02 (s, 1H), 3.87 (d, J = 8.8 Hz, 1H), 3.66 (dd, J = 8.7, 4.4 Hz, 1H), 3.54 (s, 3H), 1.37 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 202.5, 171.1, 160.0, 146.6, 145.7, 142.9, 138.6, 132.9, 129.2, 128.9, 128.7, 128.1, 127.9, 126.9, 126.8, 126.3, 126.0, 120.8, 113.2, 112.4, 80.8, 77.4, 68.0, 63.3, 59.6, 54.5, 27.2. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 11.83 min, 16.85 min. HRMS (m/z) calculated: C₃₅H₃₆NO₄ [M+H]⁺: 534.2644, found: 534.2637.</p>

 <p>4.18, 78%, d.r. 95:5</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.18 (21.4 mg, 0.102 mmol) as starting material, compound 4.18 was obtained as colorless solid (40 mg, 78%, mp 171.2 °C).</p> <p>¹H NMR (600 MHz, acetone) δ 8.32 (d, J = 2.1 Hz, 1H), 8.02 (d, J = 7.6 Hz, 2H), 7.95 (dd, J = 8.2, 1.7 Hz, 2H), 7.50 (dd, J = 16.4, 7.6 Hz, 3H), 7.42 (dt, J = 15.6, 7.5 Hz, 3H), 7.28 (t, J = 7.3 Hz, 1H), 7.19 – 7.12 (m, 3H), 7.06 (t, J = 7.7 Hz, 2H), 6.99 (t, J = 7.3 Hz, 1H), 6.86 (d, J = 7.7 Hz, 2H), 5.42 (d, J = 4.5 Hz, 1H), 4.01 (d, J = 12.1 Hz, 1H), 3.92 – 3.85 (m, 1H), 3.74 (dd, J = 8.7, 4.4 Hz, 1H), 1.34 (s, 9H). ¹³C NMR (151 MHz, acetone) δ 196.3, 171.0, 146.3, 145.4, 145.3, 142.8, 141.2, 139.4, 130.1, 128.8, 128.3, 128.1, 127.9, 127.7, 126.9, 126.8, 126.8, 126.4, 126.2, 125.1, 122.9, 80.9, 77.3, 67.6, 64.6, 59.4, 27.2. HPLC Diacel IC (5μm), Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 24.57 min, 31.53 min. HRMS (m/z) calculated: C₃₆H₃₄NO₃S [M+H]⁺: 560.2259, found: 560.2272</p>
 <p>4.19, 69%, d.r. >99:1</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.19 (19.82 mg, 0.102 mmol) as starting material, compound 4.19 was obtained as colorless solid (34 mg, 69%, mp 121.5 °C).</p> <p>¹H NMR (500 MHz, acetone) δ 7.99 (d, J = 7.6 Hz, 2H), 7.80 (d, J = 0.9 Hz, 1H), 7.42 (t, J = 7.8 Hz, 2H), 7.37 (d, J = 7.5 Hz, 2H), 7.29 (t, J = 7.3 Hz, 1H), 7.23 – 7.11 (m, 4H), 7.07 (t, J = 7.6 Hz, 2H), 7.00 (t, J = 7.3 Hz, 1H), 6.88 (dd, J = 6.5, 2.8 Hz, 2H), 6.58 (dd, J = 3.6, 1.6 Hz, 1H), 5.21 (d, J = 4.3 Hz, 1H), 3.91 (s, 1H), 3.86 – 3.73 (m, 2H), 1.35 (s, 9H). ¹³C NMR (126 MHz, acetone) δ 192.5, 173.8, 156.2, 149.9, 149.0, 147.3, 144.00, 131.4, 130.9, 130.7, 130.4, 129.7, 129.6, 129.4, 128.88, 128.86, 120.2, 115.4, 83.5, 79.6, 69.9, 66.6, 60.7, 29.9. HPLC: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 14.52 min, 16.13 min. HRMS (m/z) calculated: C₃₂H₃₂NO₄ [M+H]⁺: 494.2331, found: 494.2321.</p>
 <p>4.20, 96%, d.r. 99:1</p>	<p>Following Procedure D using α,β-unsaturated ketone 2.20 (23.8 mg, 0.102 mmol) compound 4.20 was obtained as clear oil (53 mg, 96%).</p> <p>¹H NMR (400 MHz, Acetone) δ 8.41 (s, 1H), 8.04 – 7.99 (m, 2H), 7.90 (dd, J = 16.4, 8.5 Hz, 3H), 7.73 (dd, J = 8.7, 1.7 Hz, 1H), 7.65 – 7.59 (m, 1H), 7.57 – 7.52 (m, 1H), 7.43 – 7.36 (m, 4H), 7.26 (t, J = 7.3 Hz, 1H), 7.17 (dd, J = 5.0, 1.9 Hz, 3H), 7.05 – 6.95 (m, 3H), 6.91 (dd, J = 6.4, 3.1 Hz, 2H), 5.60 (d, J = 4.8 Hz, 1H), 3.95 (d, J = 8.7 Hz, 1H), 3.79 (dd, J = 8.7, 4.8 Hz, 1H), 1.35 (s, 9H). ¹³C NMR (101 MHz, Acetone) δ 202.0, 171.2, 146.6, 145.6, 141.3, 135.9, 135.5, 132.6, 129.8, 129.6, 128.8, 128.6, 128.5, 128.4, 128.2, 127.9, 127.7, 126.9, 126.9, 126.9, 126.8, 126.5, 126.1, 123.9, 80.8, 77.3, 67.8, 63.6, 59.2, 27.3. HPLC: Diacel IC (5μm), Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 34.07 min, 36.73 min. HRMS (m/z) calculated: C₃₈H₃₅NO₃Na [M+Na]⁺: 576.2515, found: 576.2512.</p>

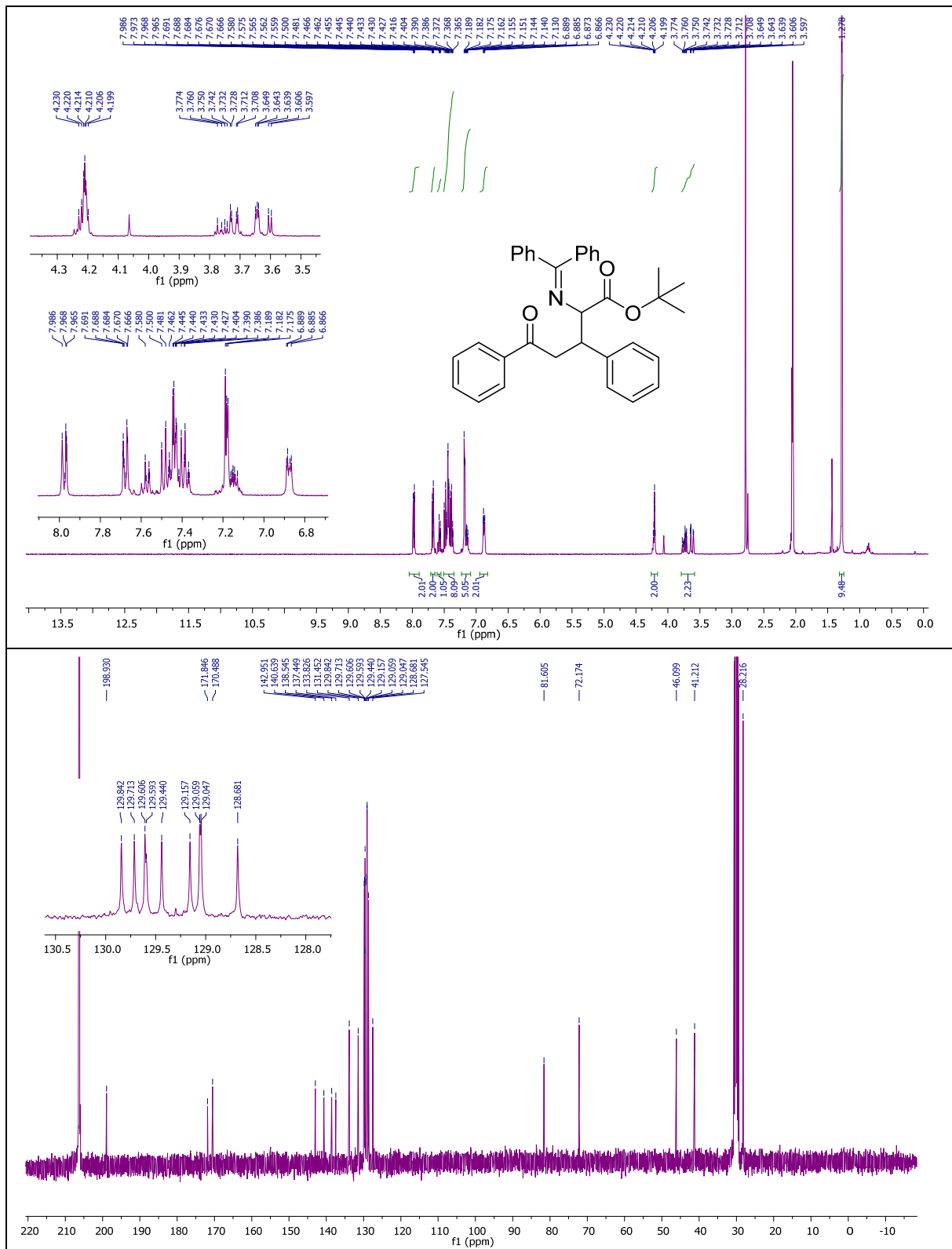


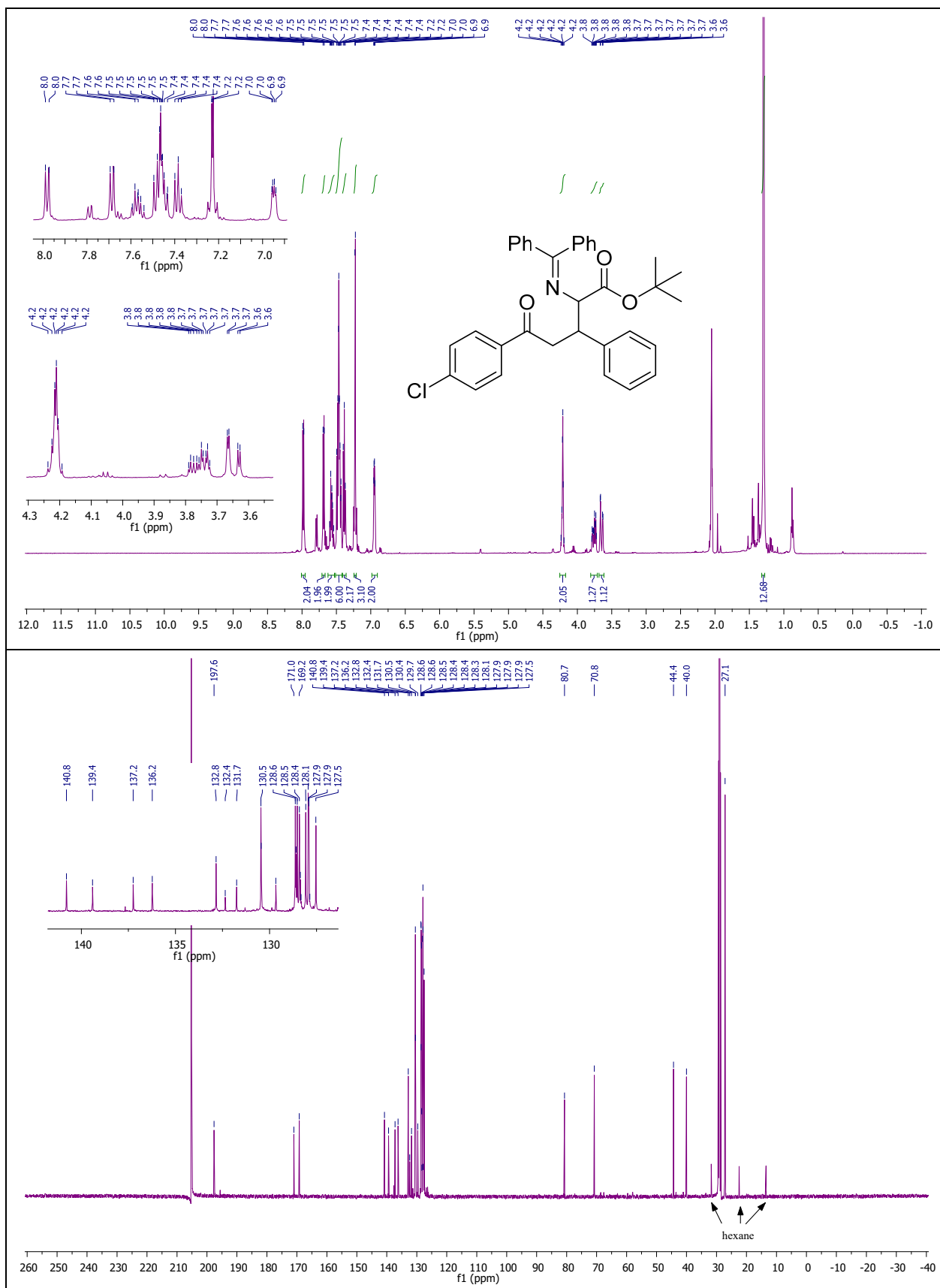
4.21, 99%, d.r. >99:1

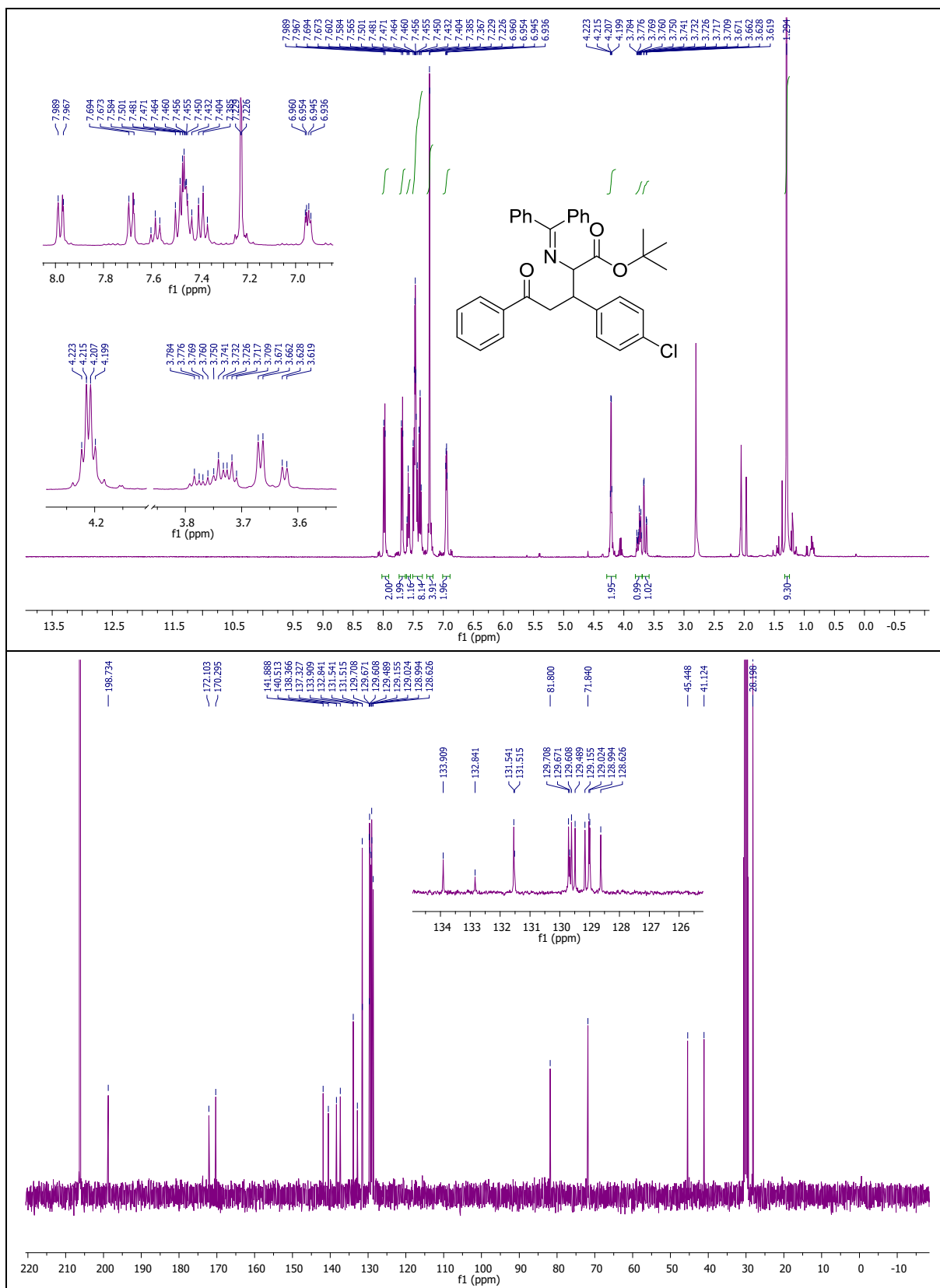
Following Procedure D using α,β -unsaturated ketone **2.21** (23.8 mg, 0.102 mmol) compound **4.21** was obtained as clear oil (55 mg, 99%).

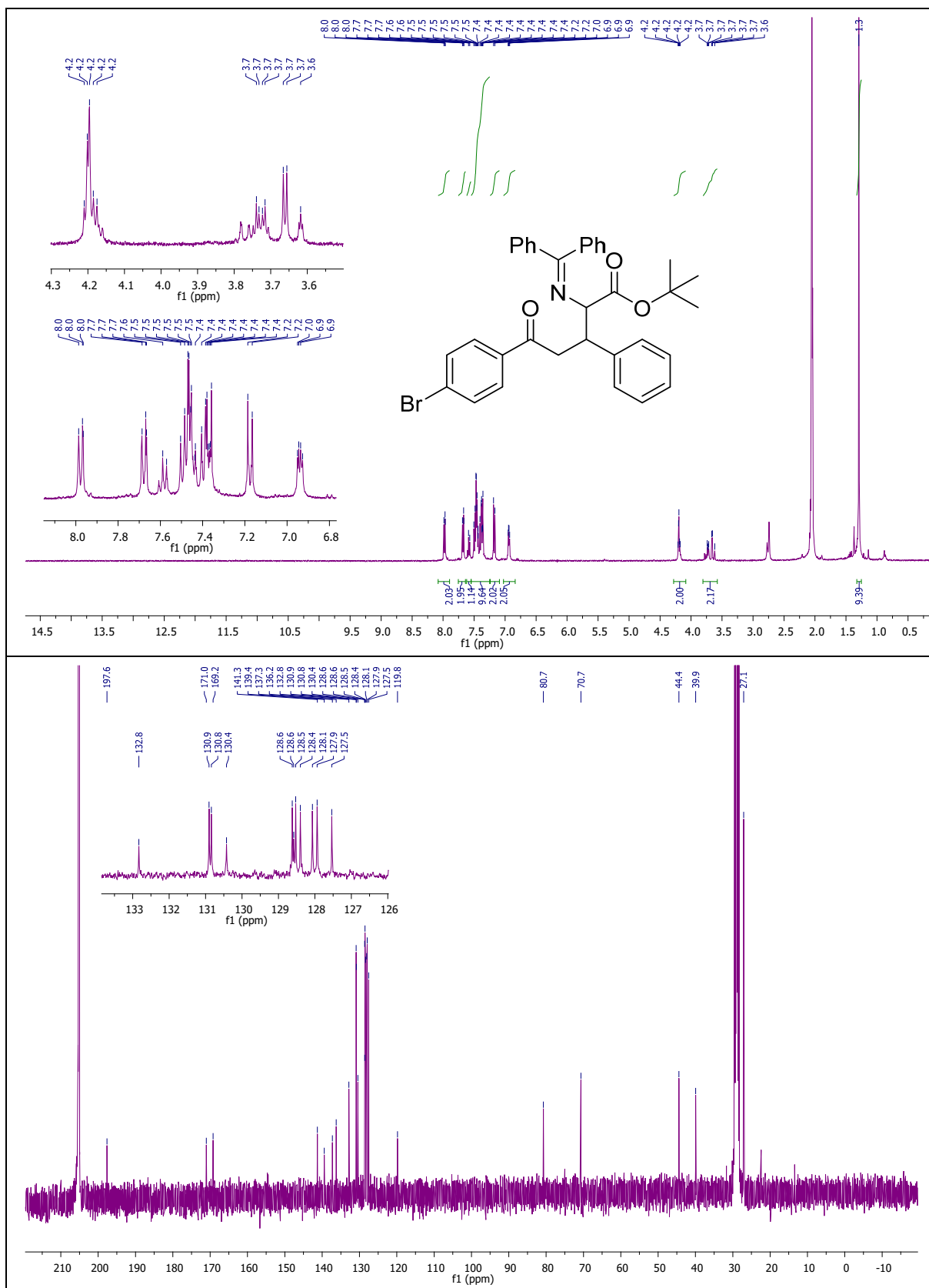
$^1\text{H NMR}$ (400 MHz, Acetone) δ 7.93 (d, J = 7.3 Hz, 2H), 7.87 (d, J = 7.6 Hz, 1H), 7.81 (d, J = 8.7 Hz, 1H), 7.75 (d, J = 8.2 Hz, 1H), 7.62 (dd, J = 8.3, 1.2 Hz, 2H), 7.49 – 7.36 (m, 4H), 7.33 – 7.22 (m, 1H), 7.04 – 6.94 (m, 3H), 6.90 (d, J = 6.7 Hz, 1H), 5.46 (d, J = 4.9 Hz, 1H), 4.85 – 4.72 (m, 1H), 4.11 (d, J = 7.1 Hz, 2H), 1.13 (s, 9H). **$^{13}\text{C NMR}$** (101 MHz, Acetone) δ 204.0, 172.3, 147.4, 145.4, 139.9, 138.5, 134.9, 133.8, 133.2, 129.9, 129.7, 129.5, 129.0, 128.7, 128.3, 128.2, 128.1, 127.7, 127.3, 127.0, 126.5, 126.4, 125.5, 124.0, 81.7, 78.6, 68.4, 64.8, 53.2, 28.0. **HPLC**: Kromasil OD-H, Hexane:2-Propanol (98:2), 0.5 mL/min, 254 nm; retention time: 11.06 min, 11.58 min. **HRMS** (m/z) calculated: $\text{C}_{38}\text{H}_{35}\text{NO}_3\text{Na}$ [$\text{M} + \text{Na}$] $^+$: 576.2515, found: 576.2494.

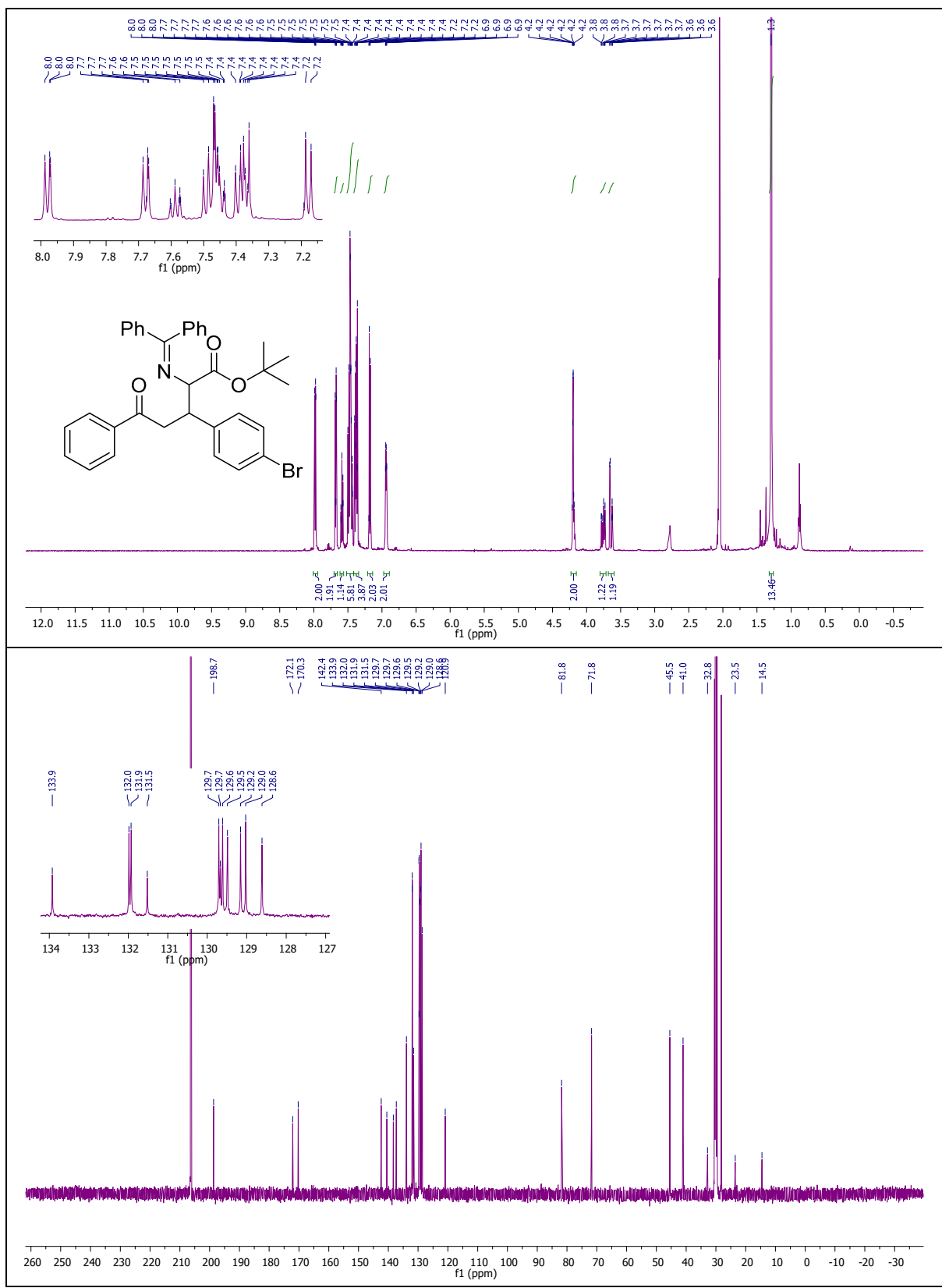
3. Copies of ^1H and ^{13}C NMR spectra

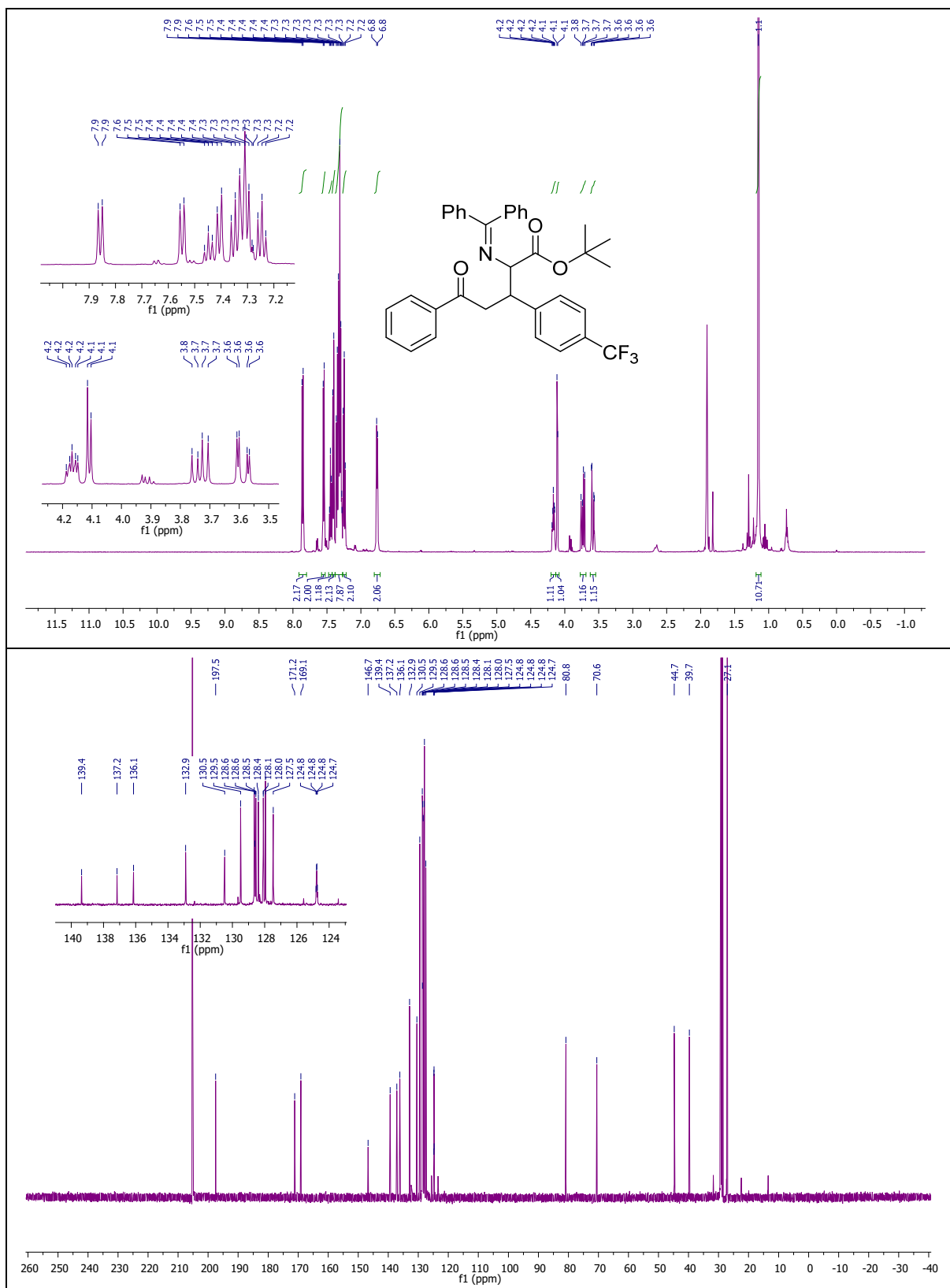


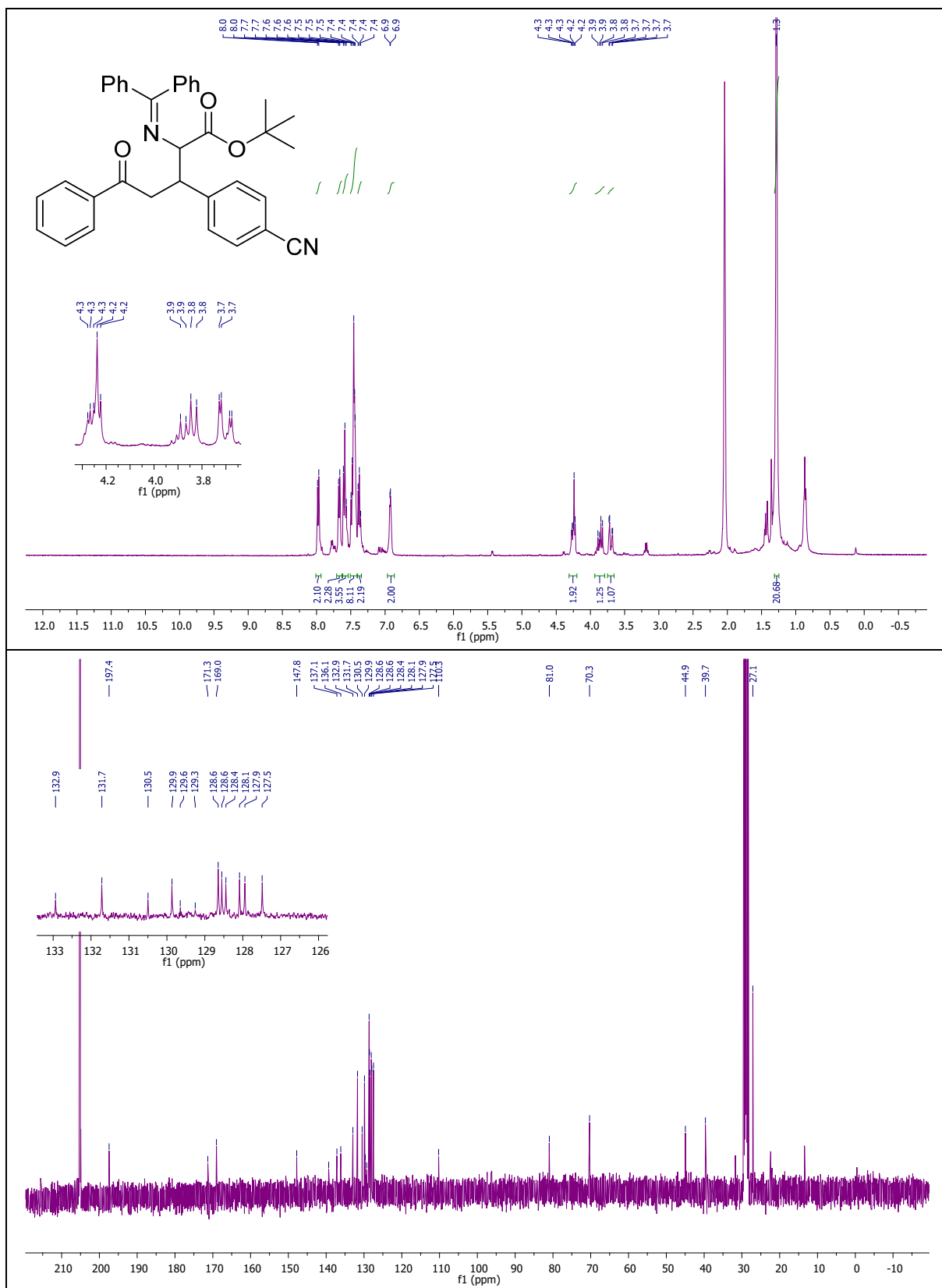


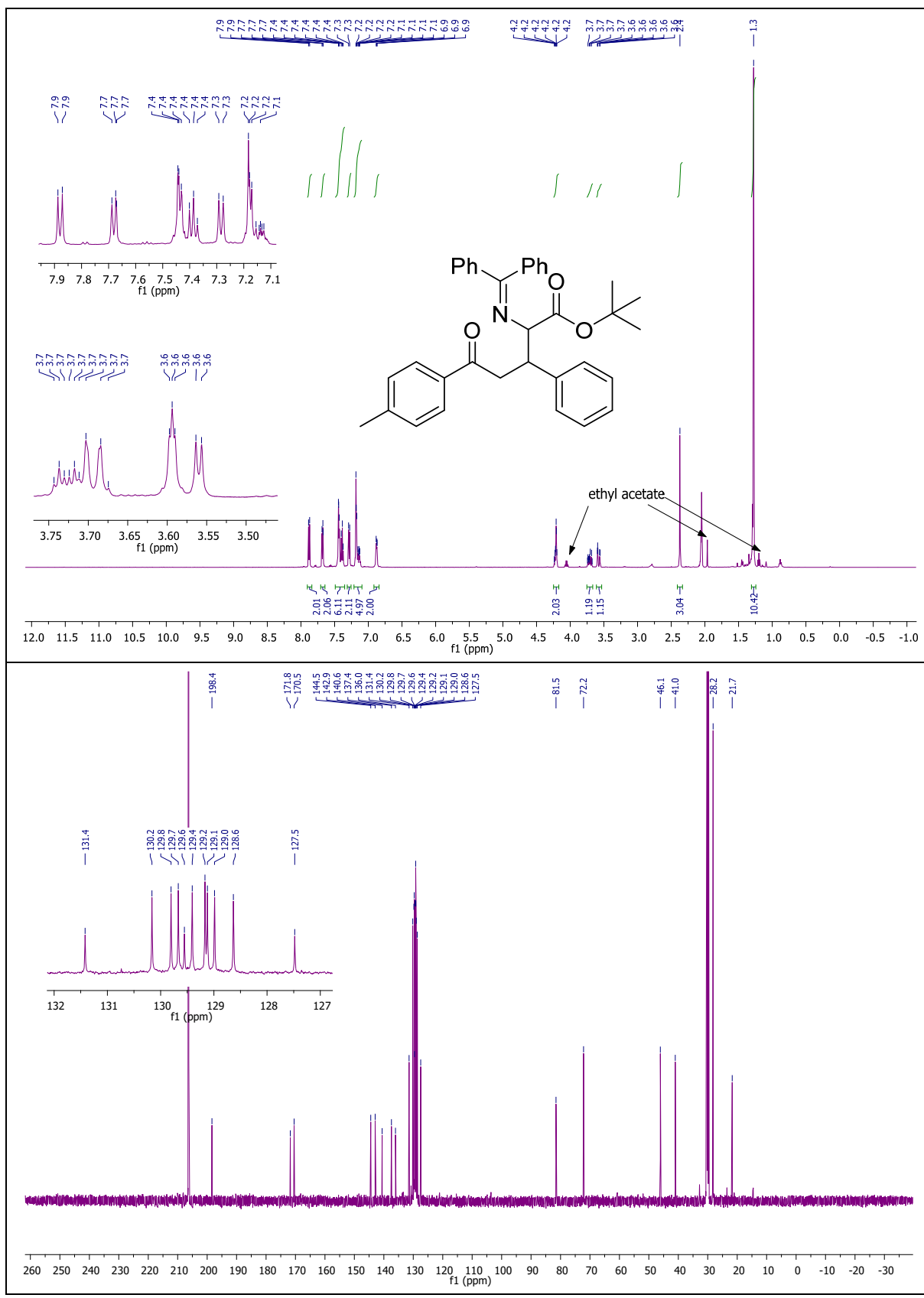


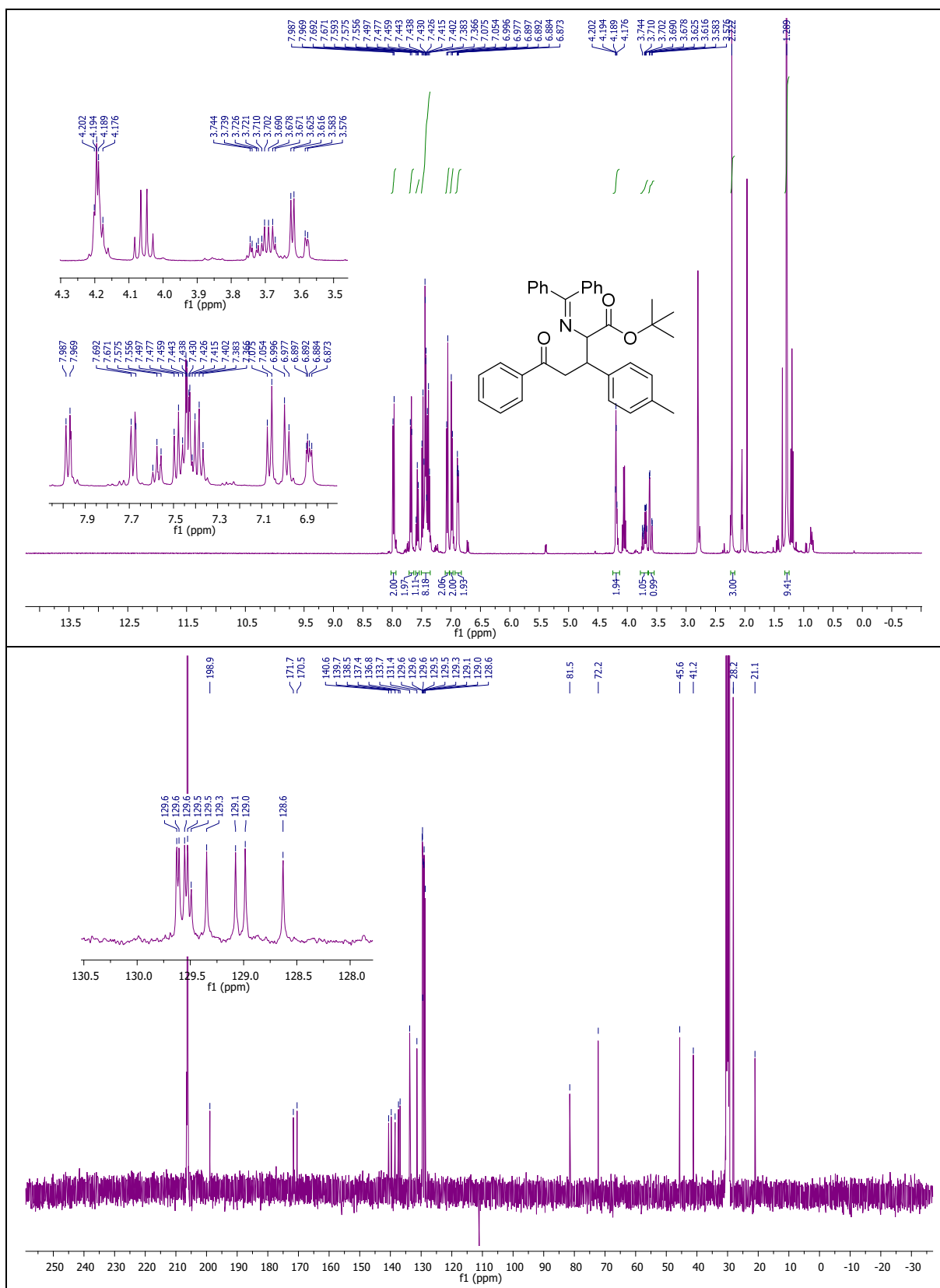


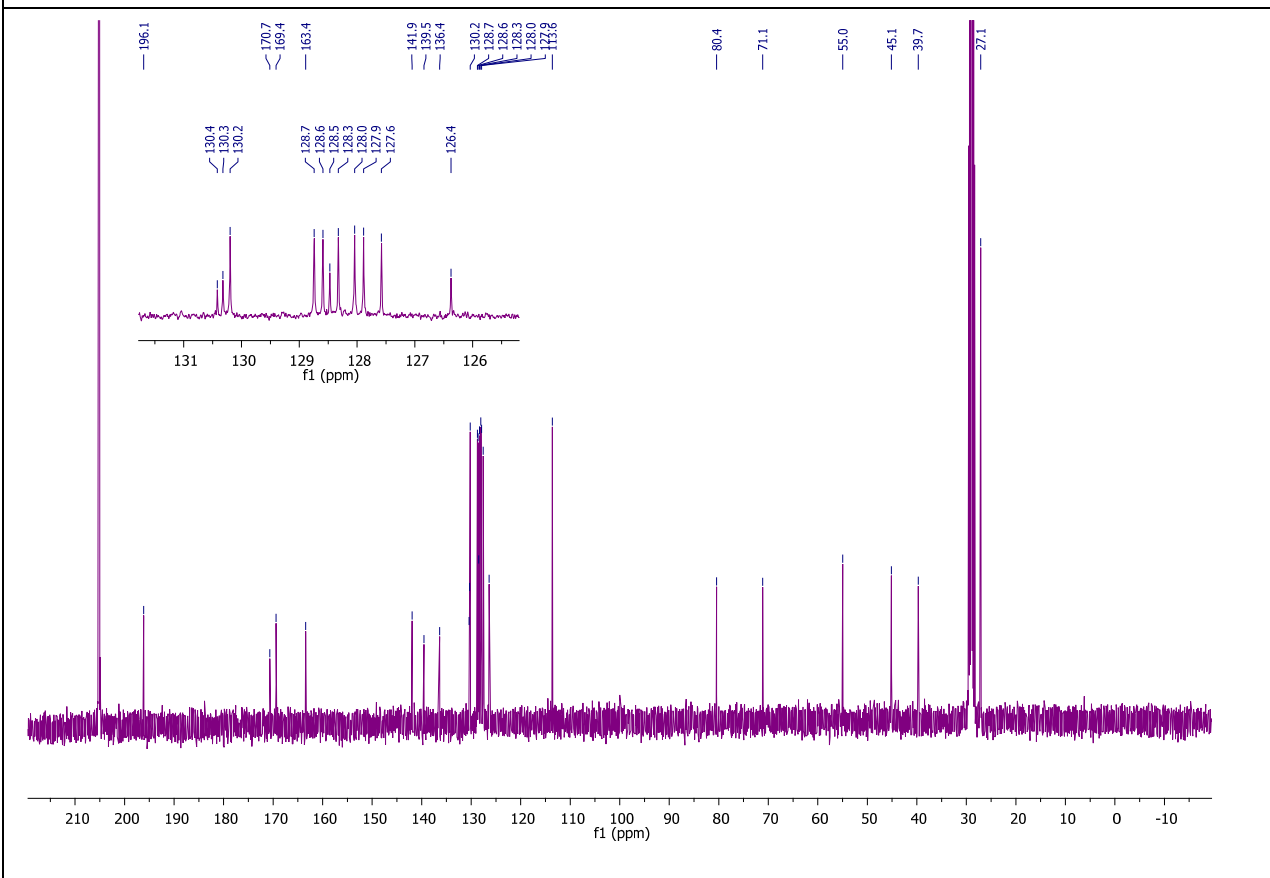
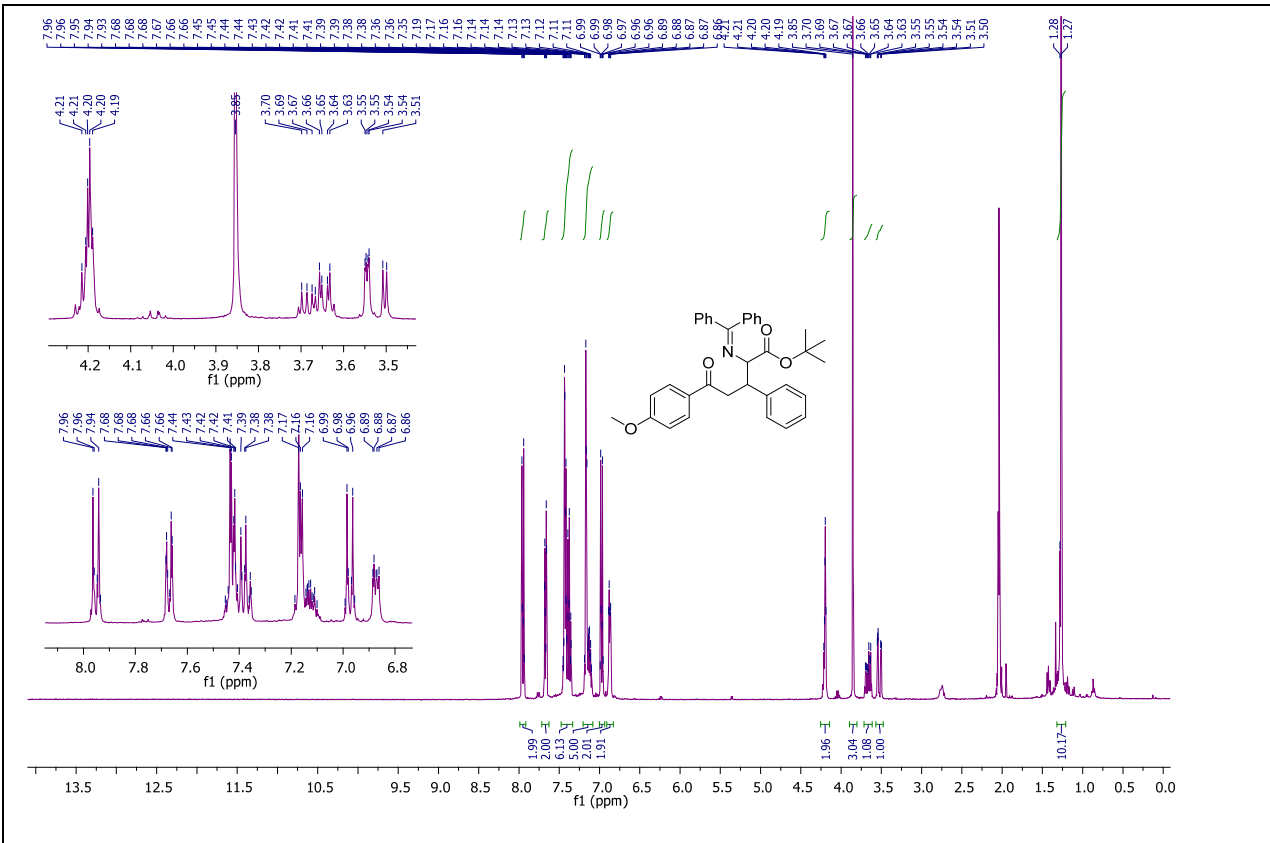


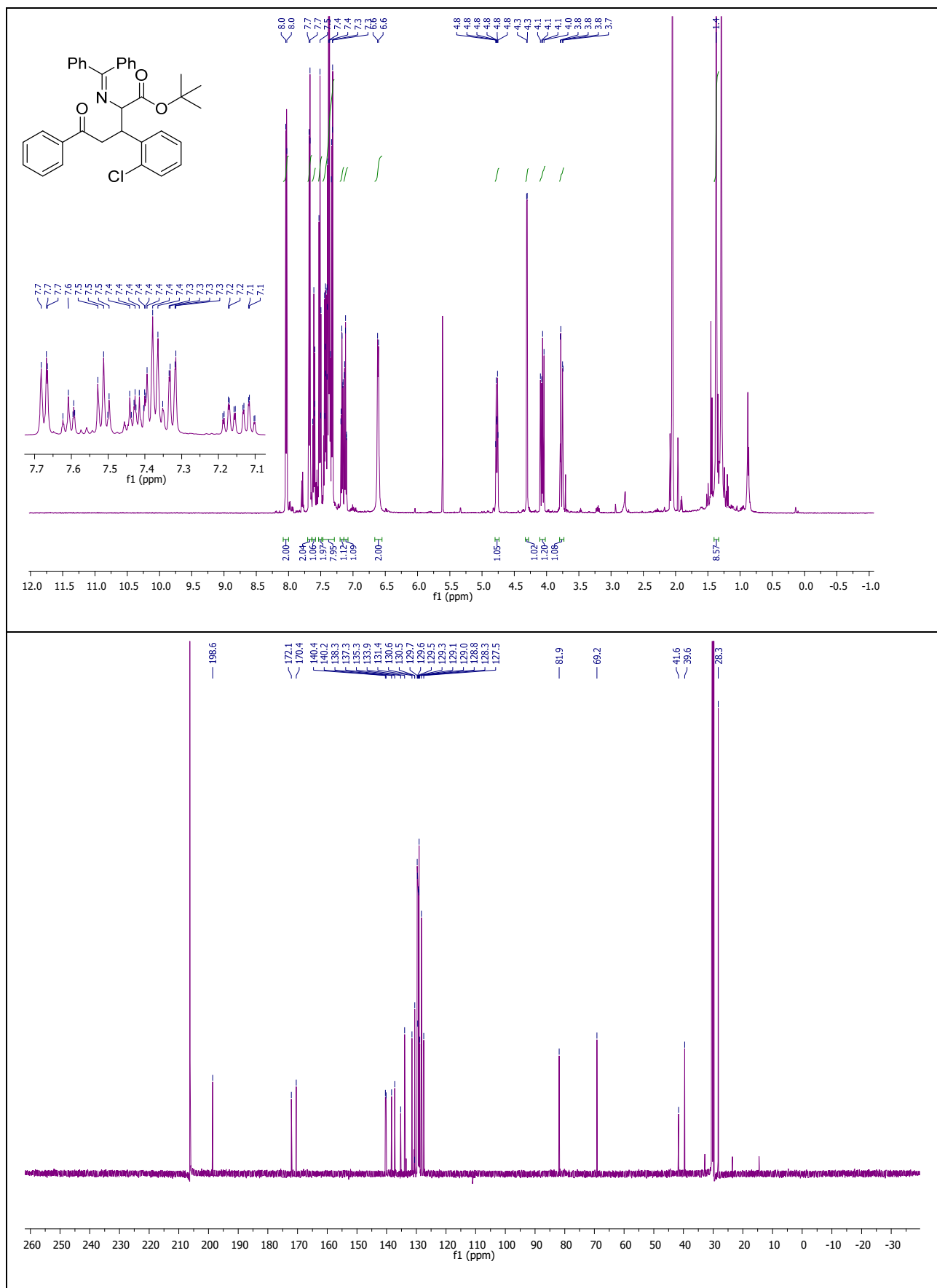


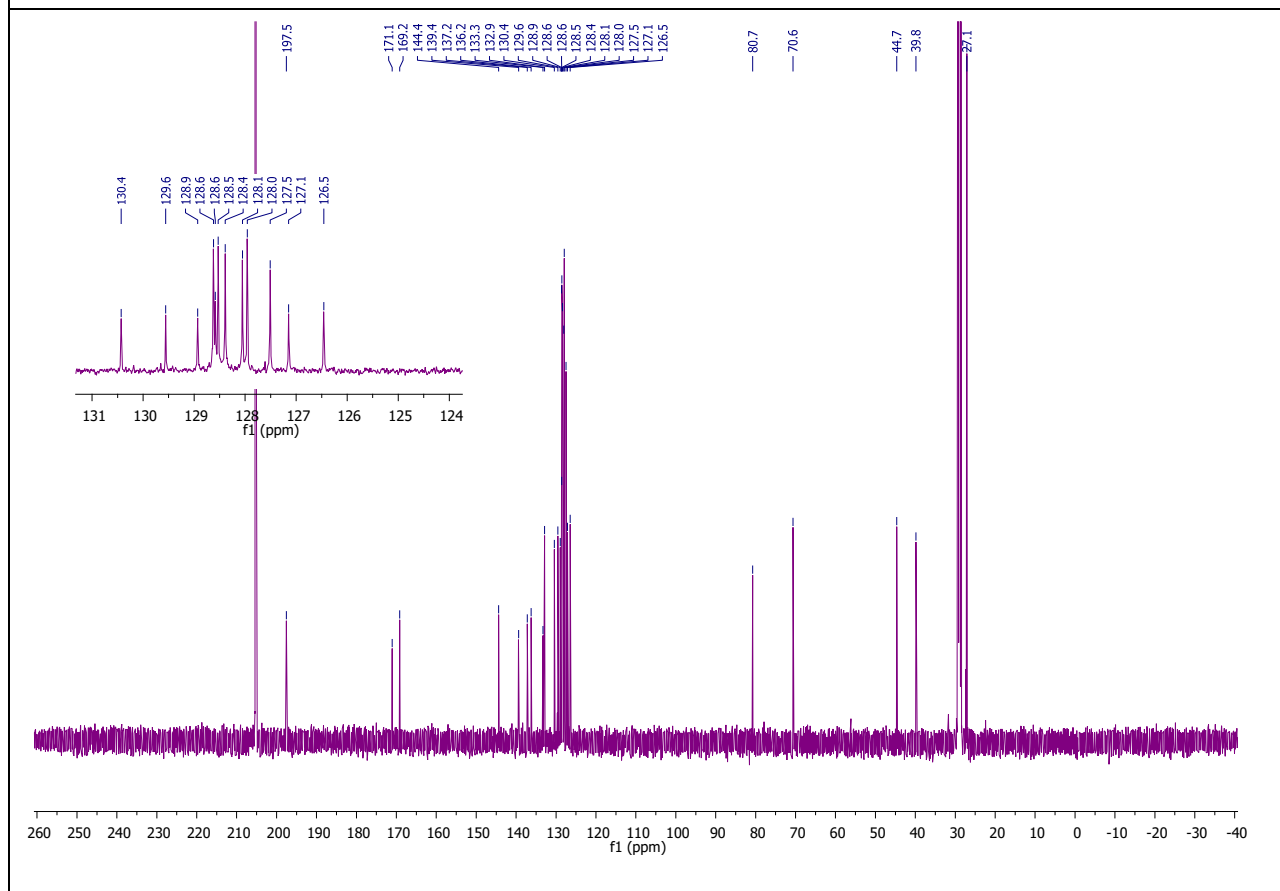
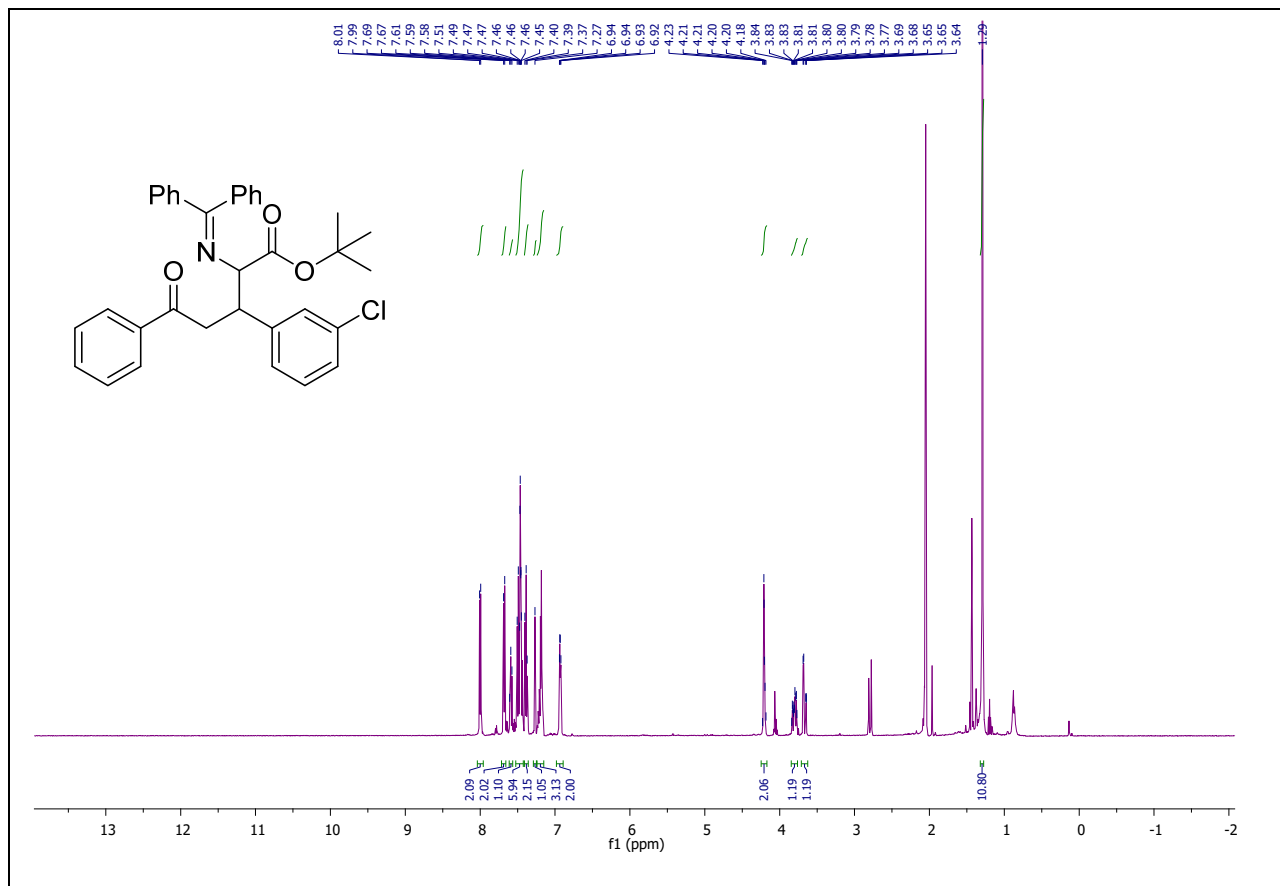


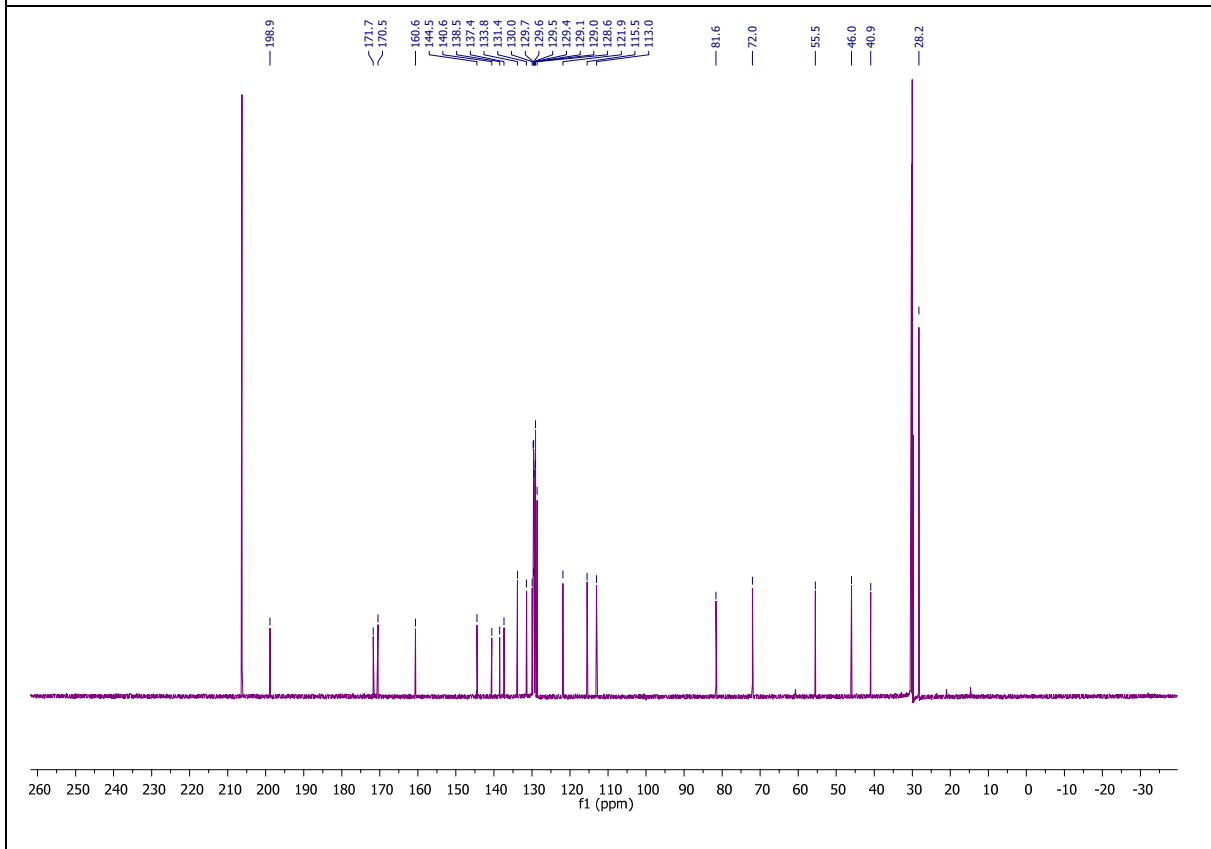
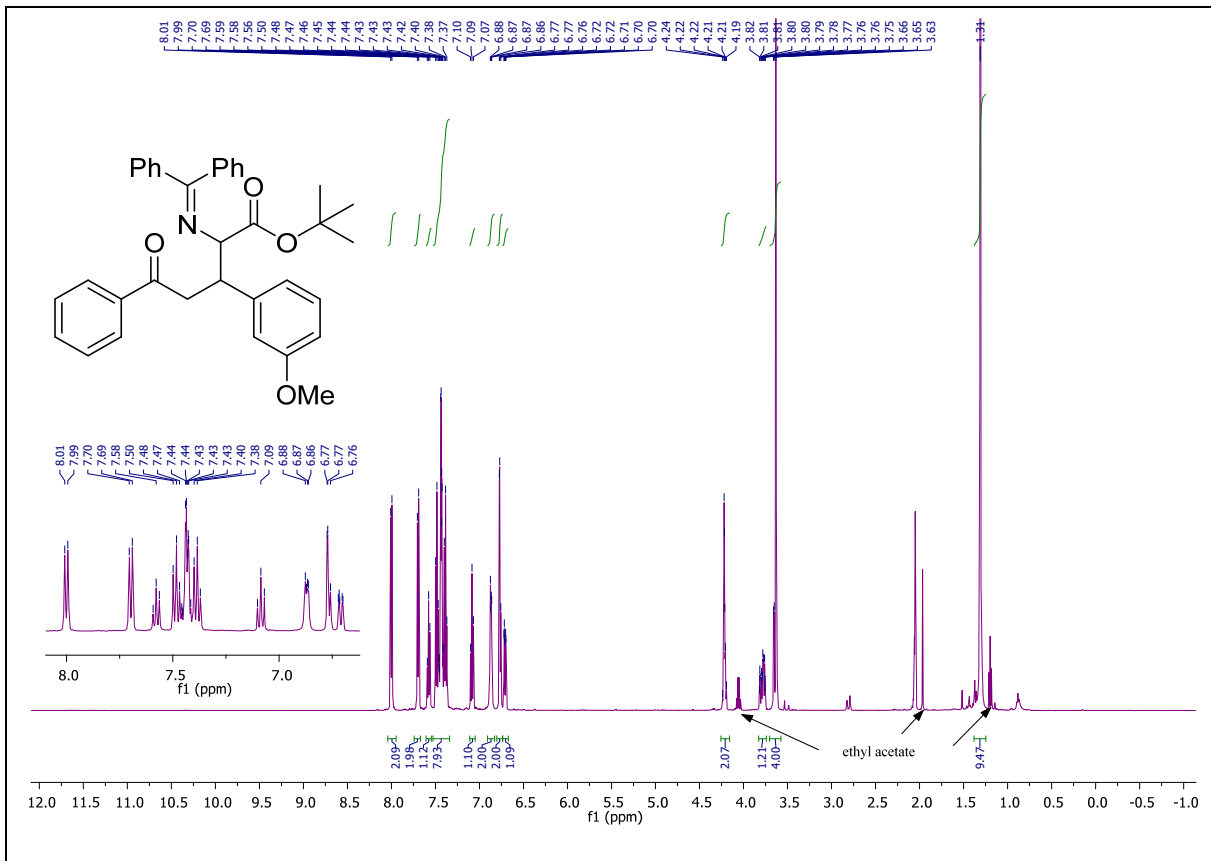


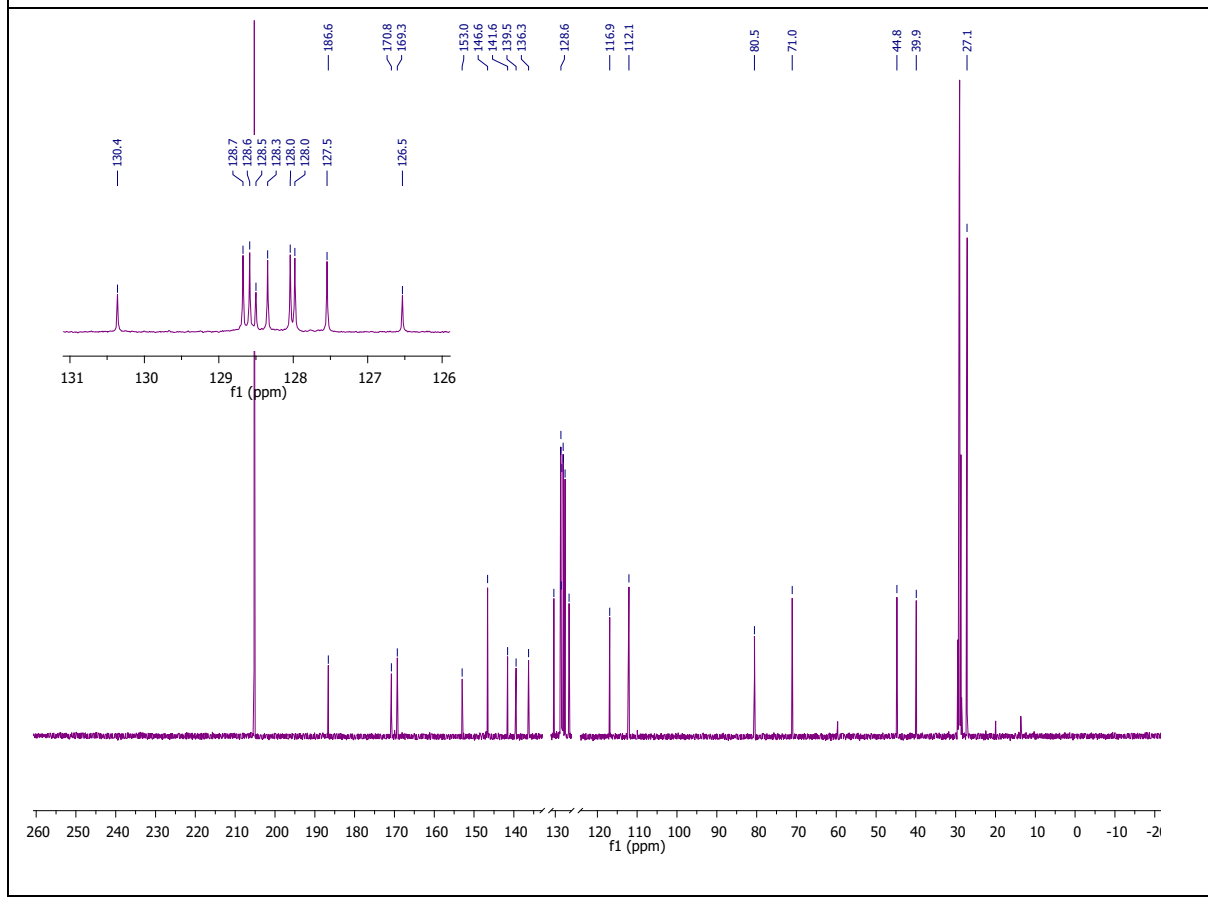
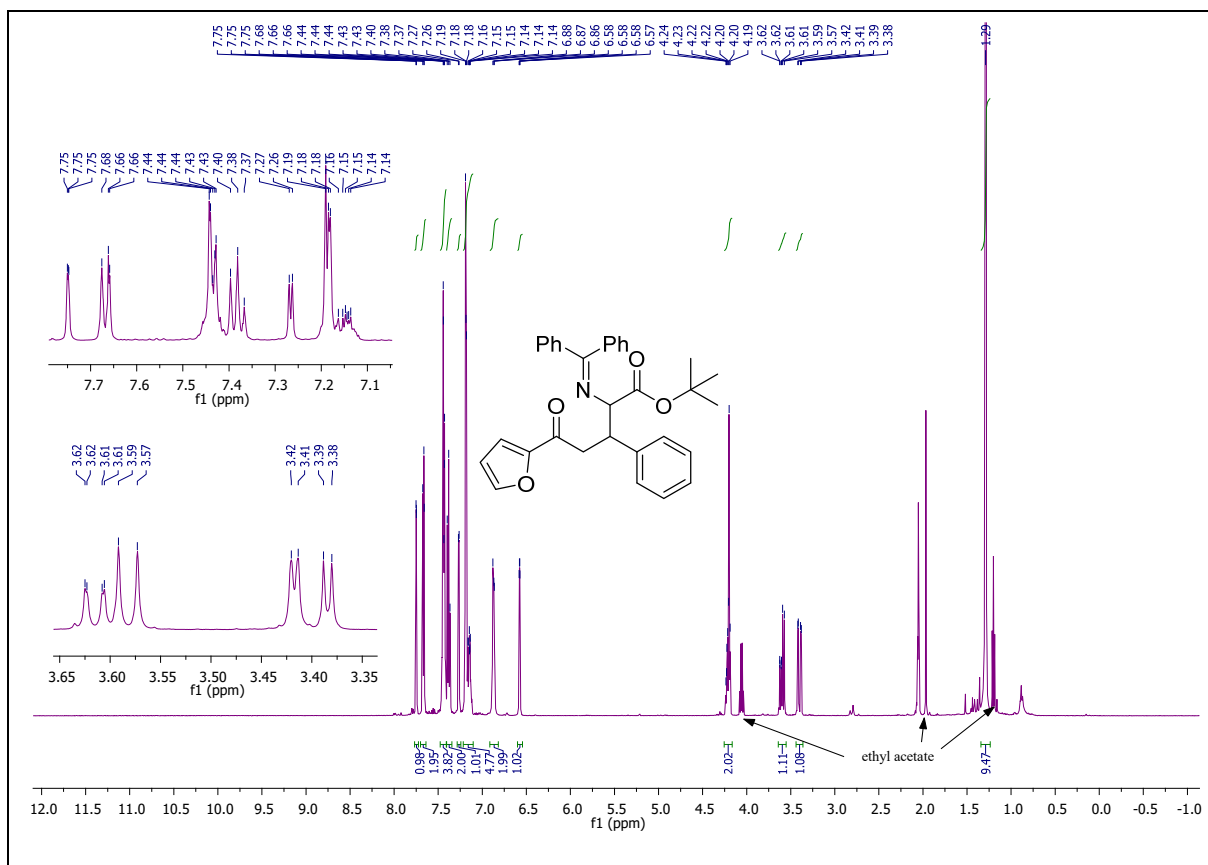


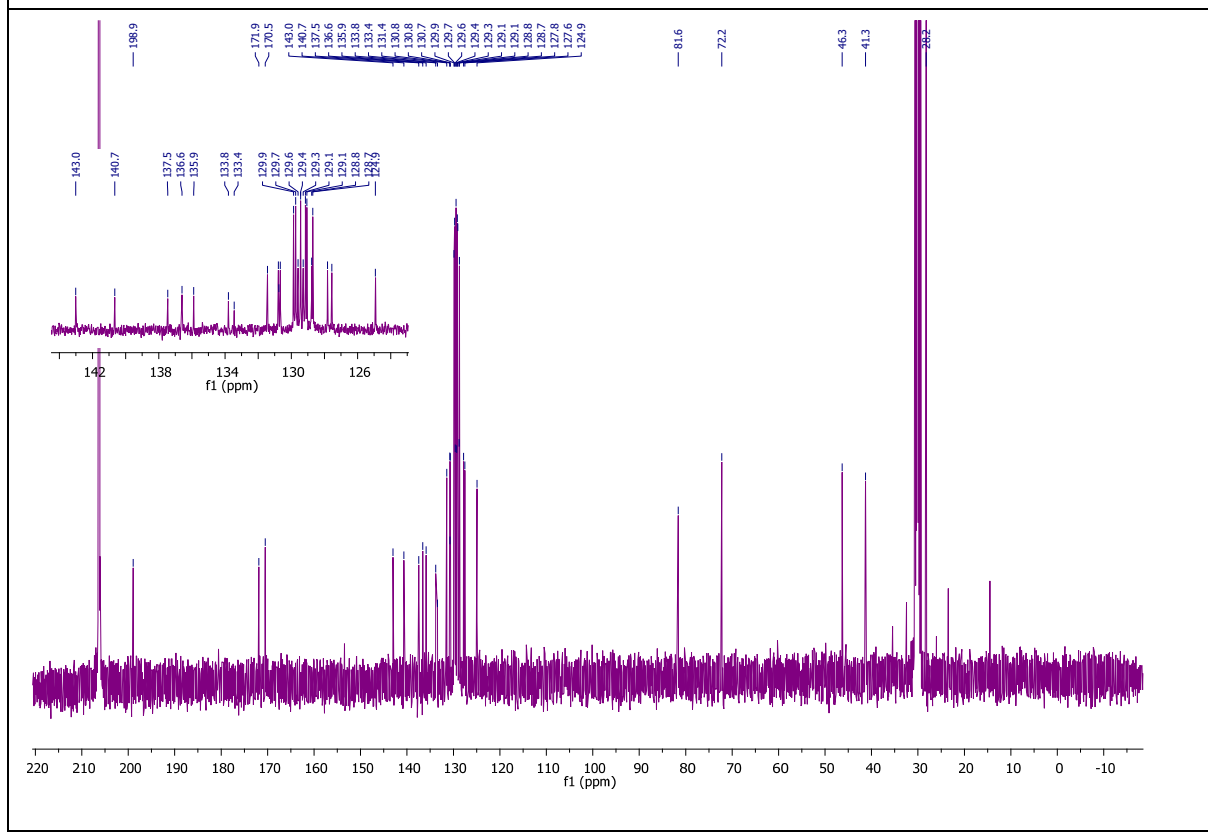
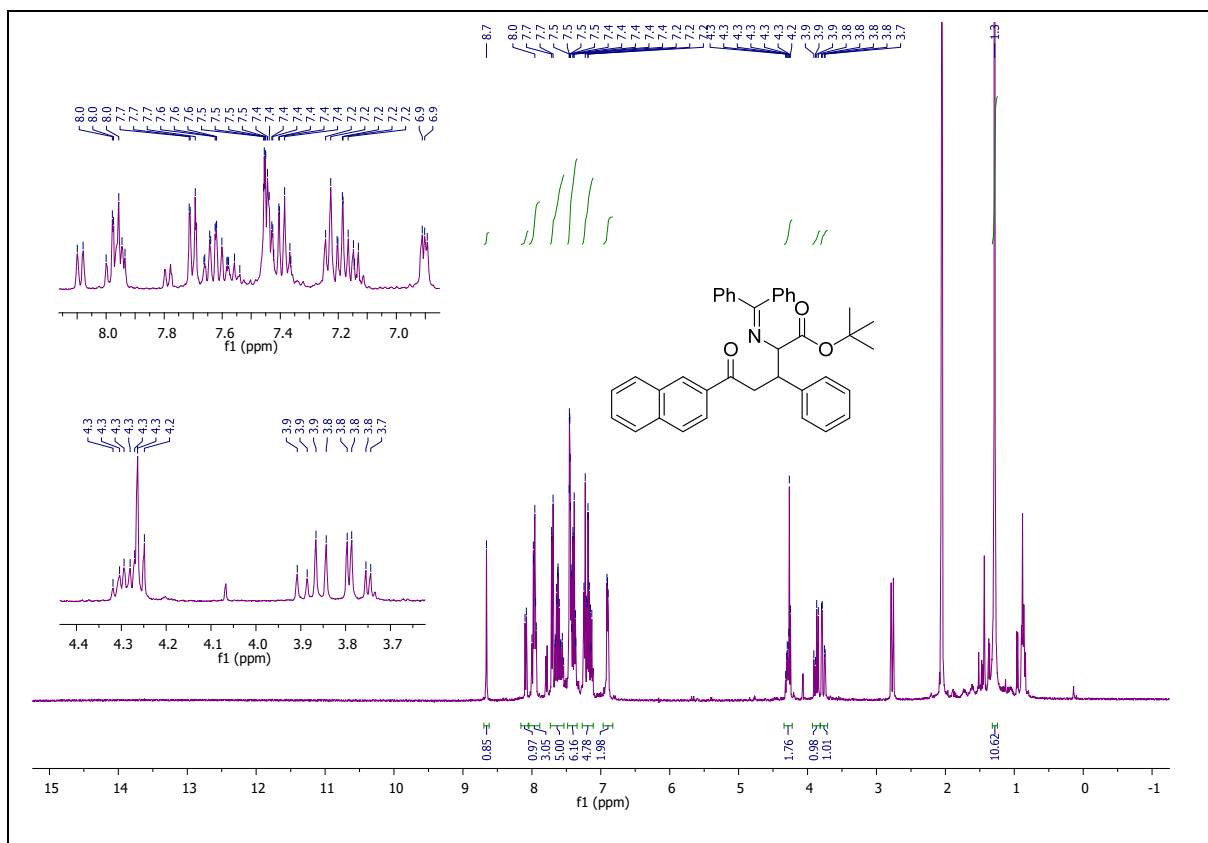


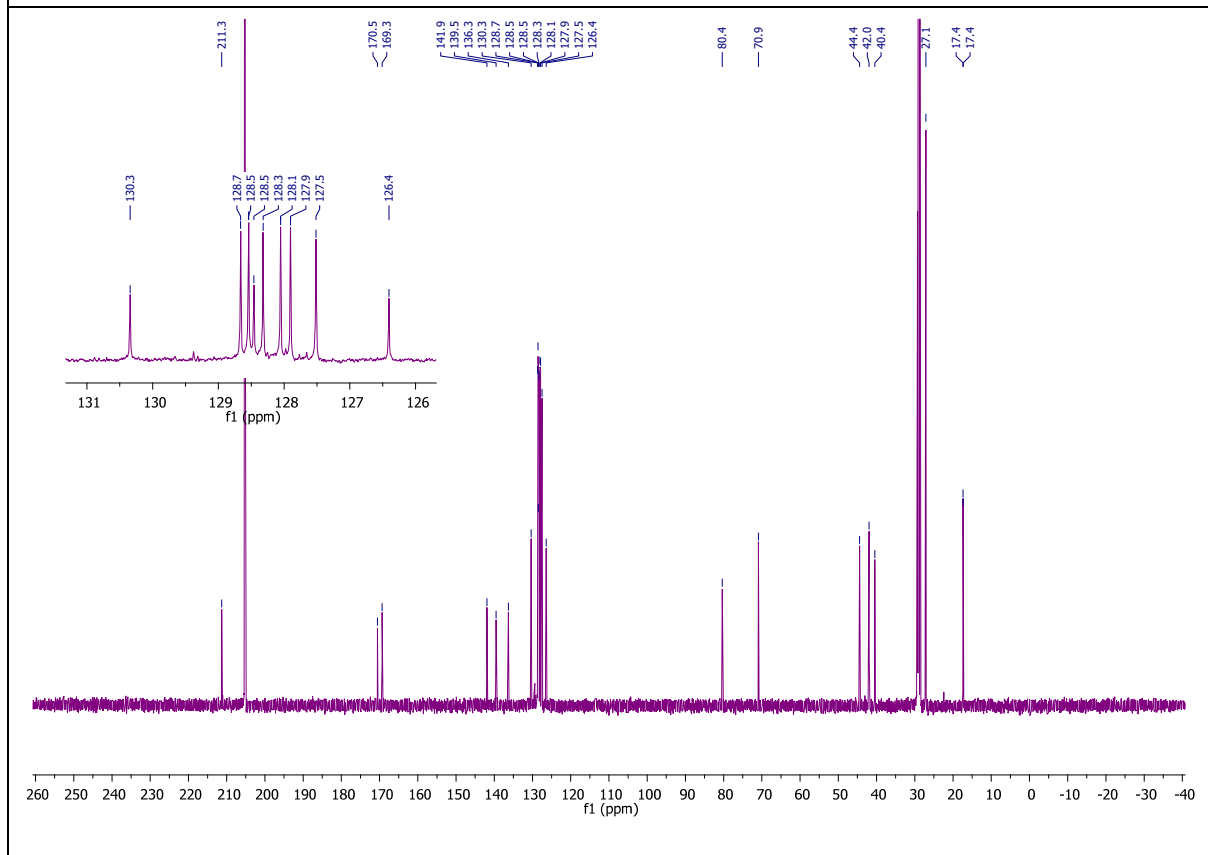
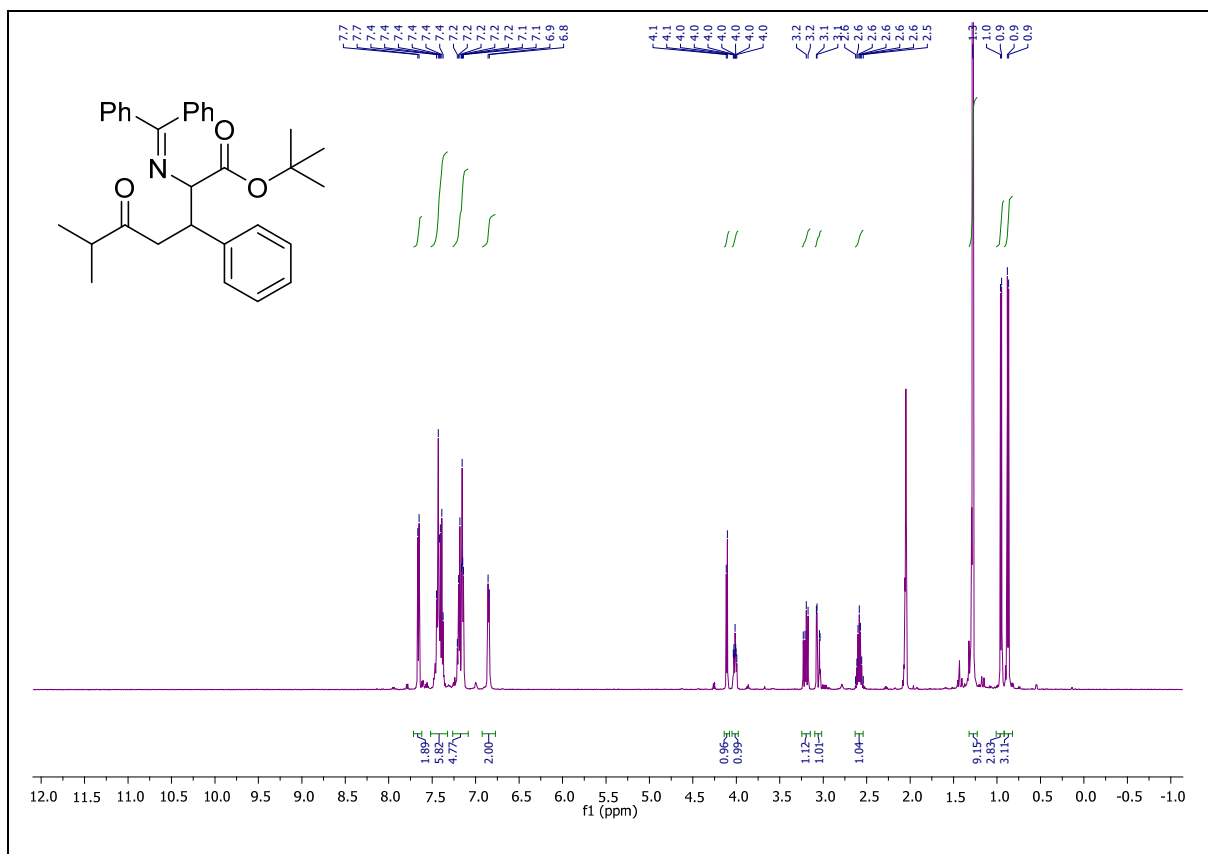


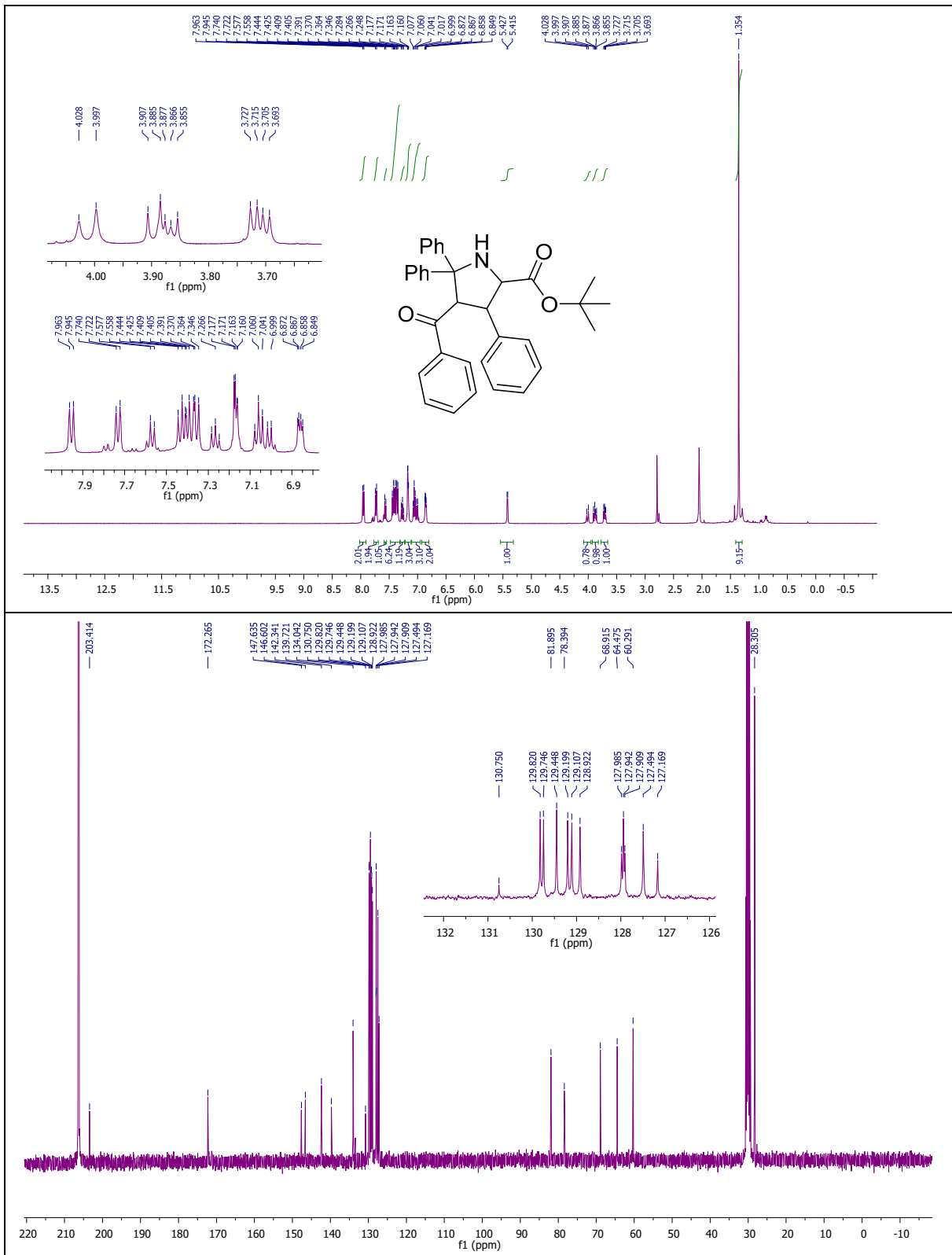


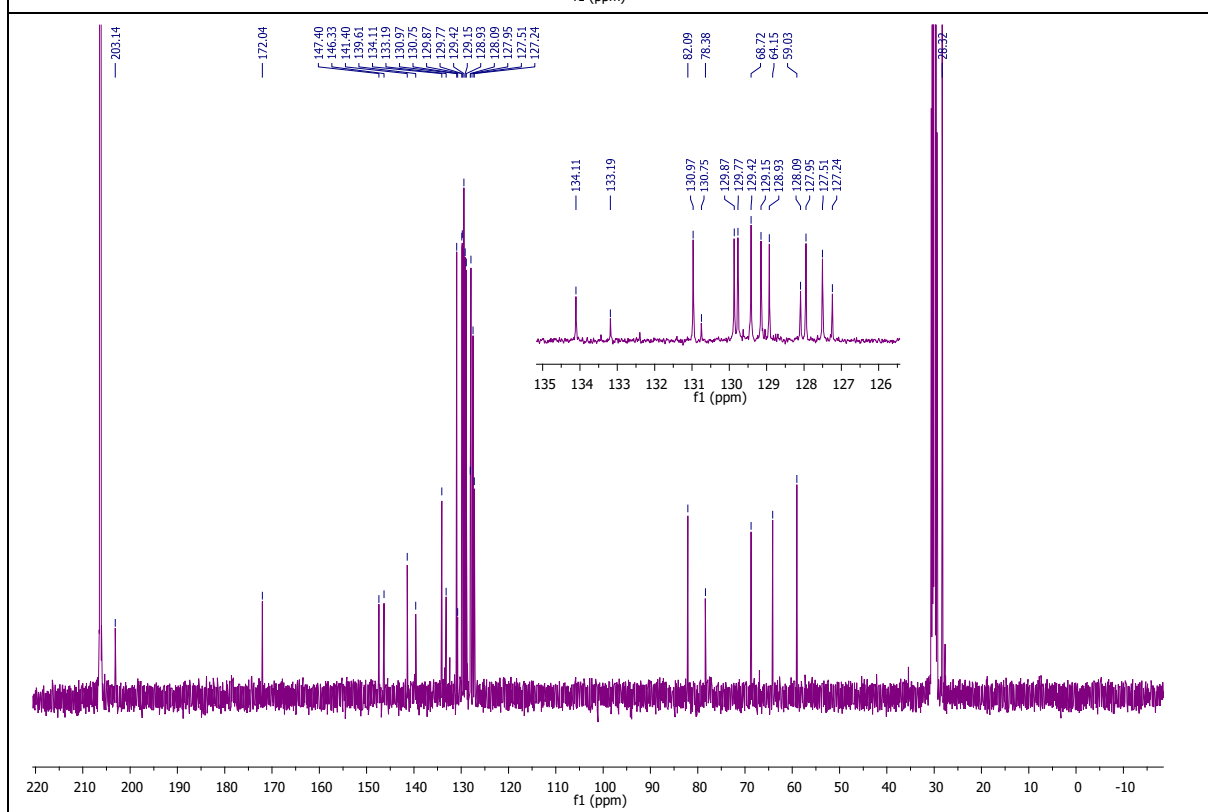
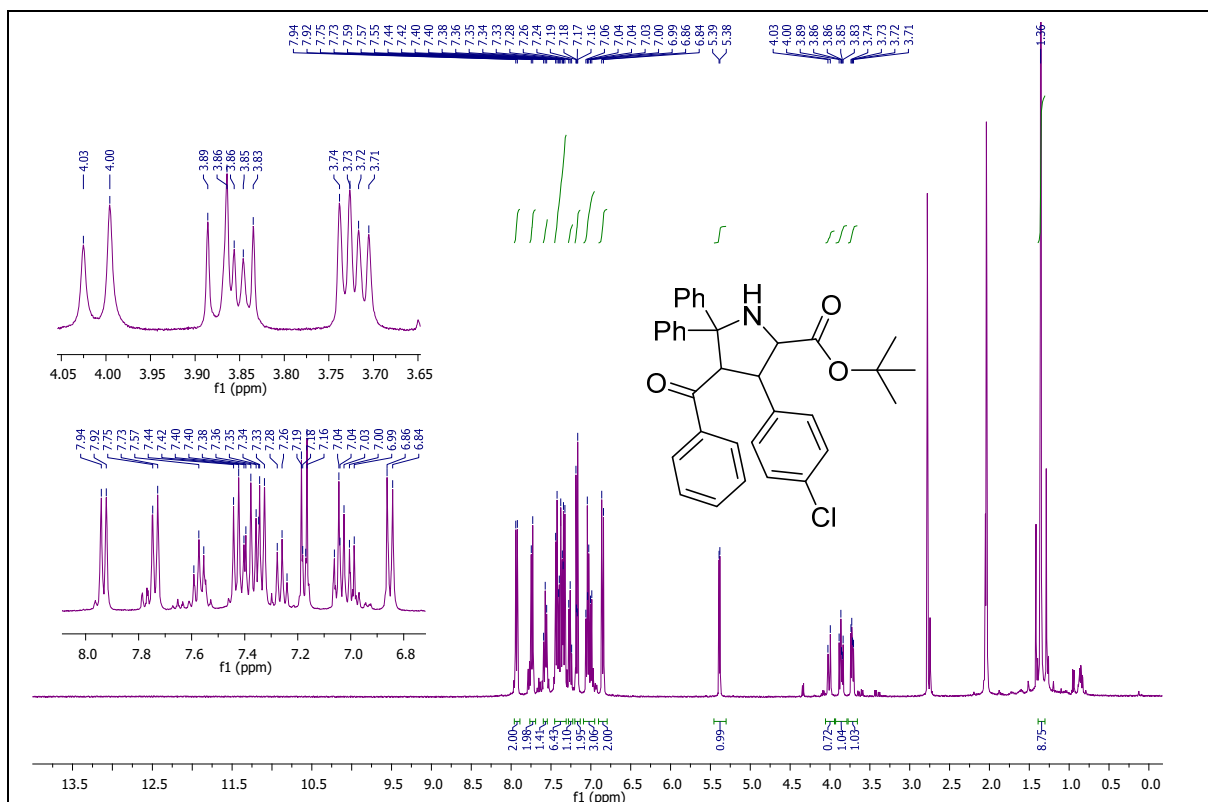


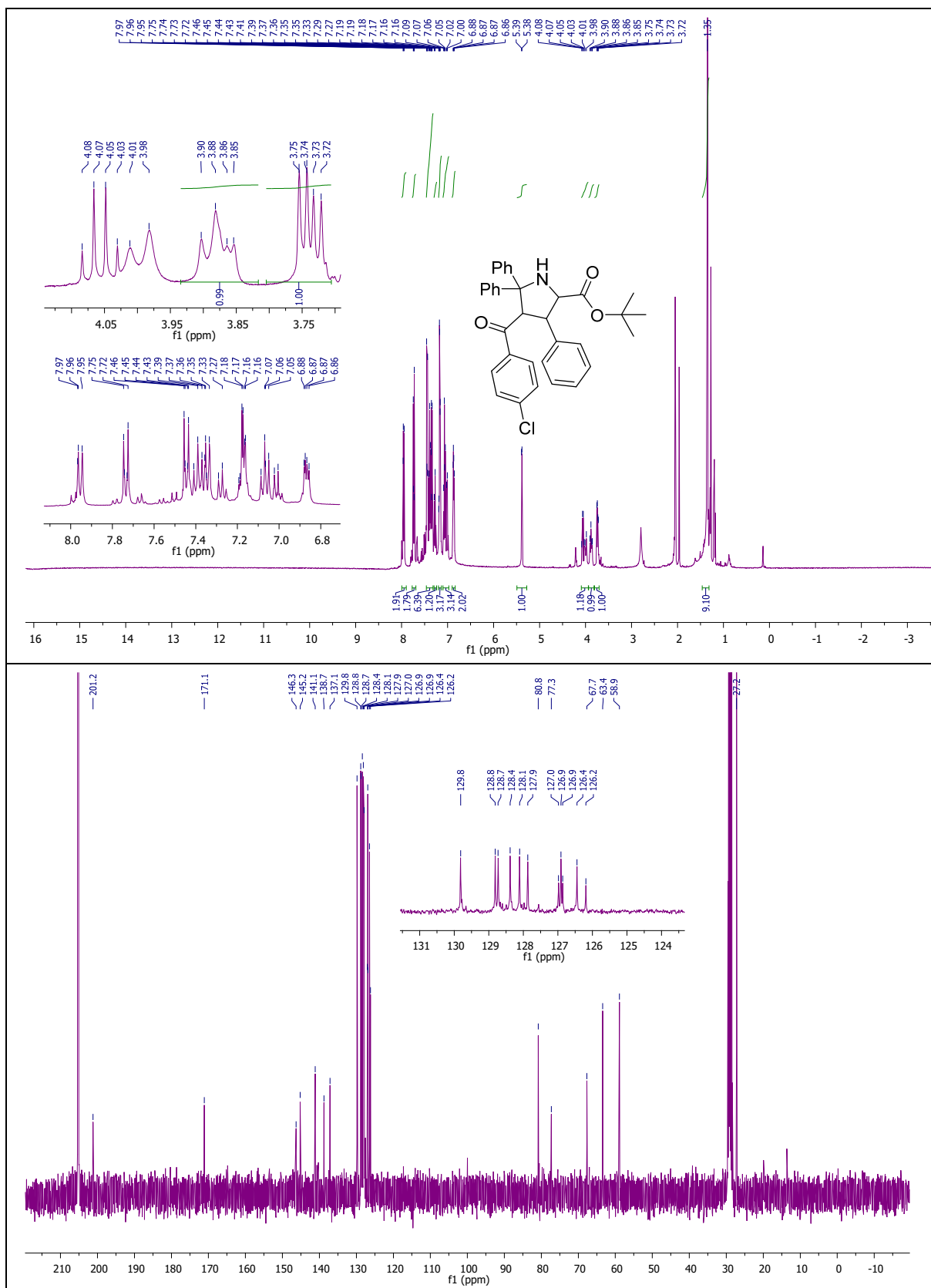


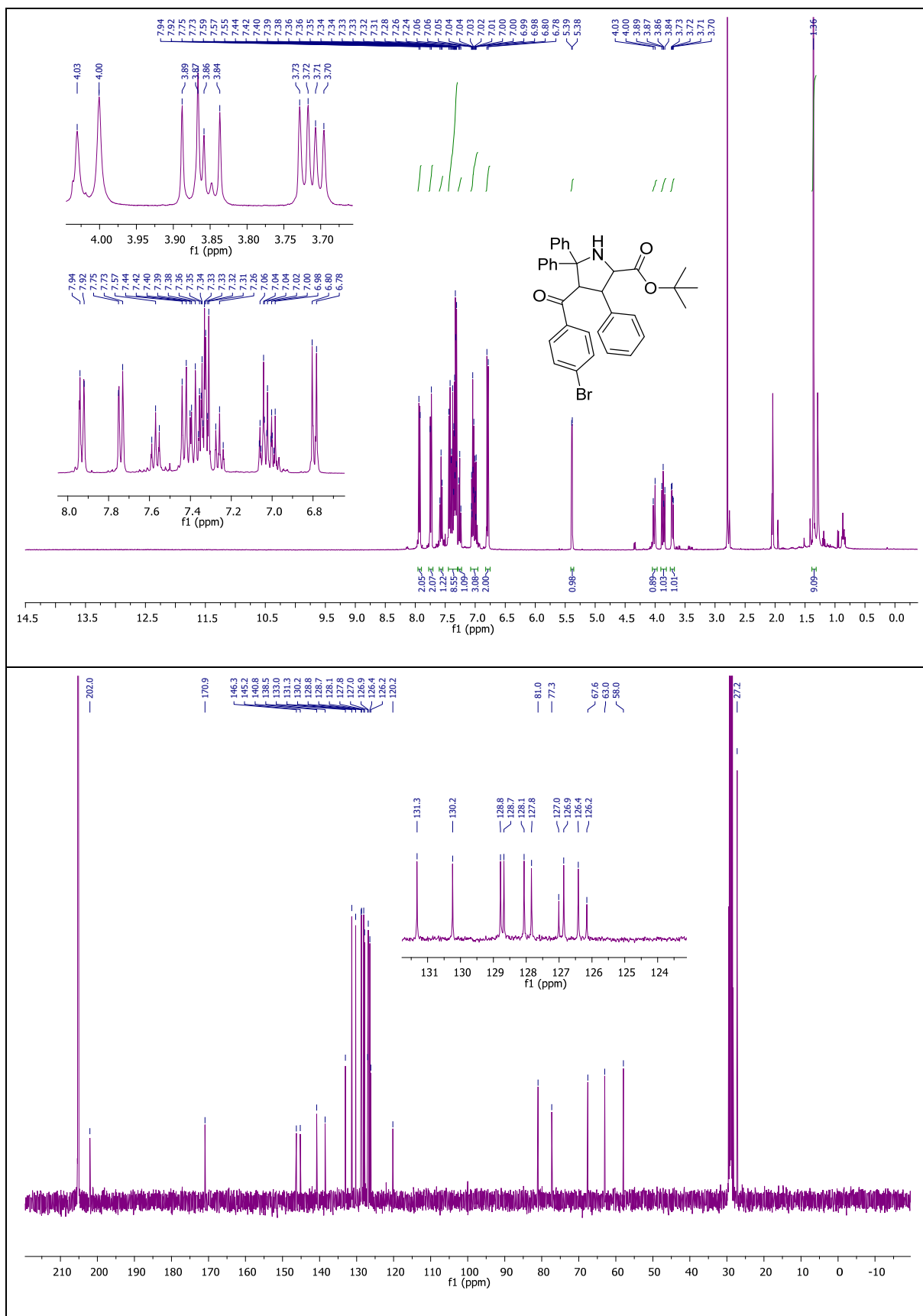


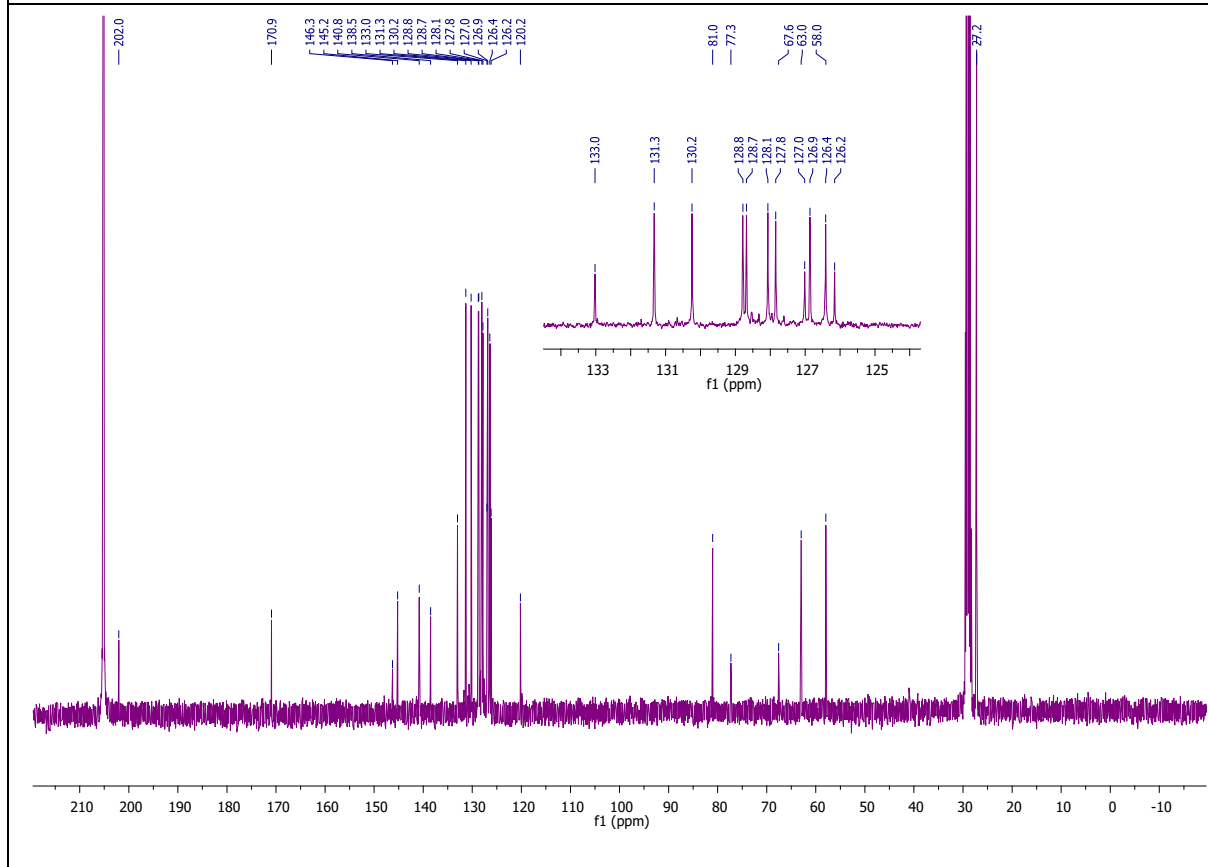
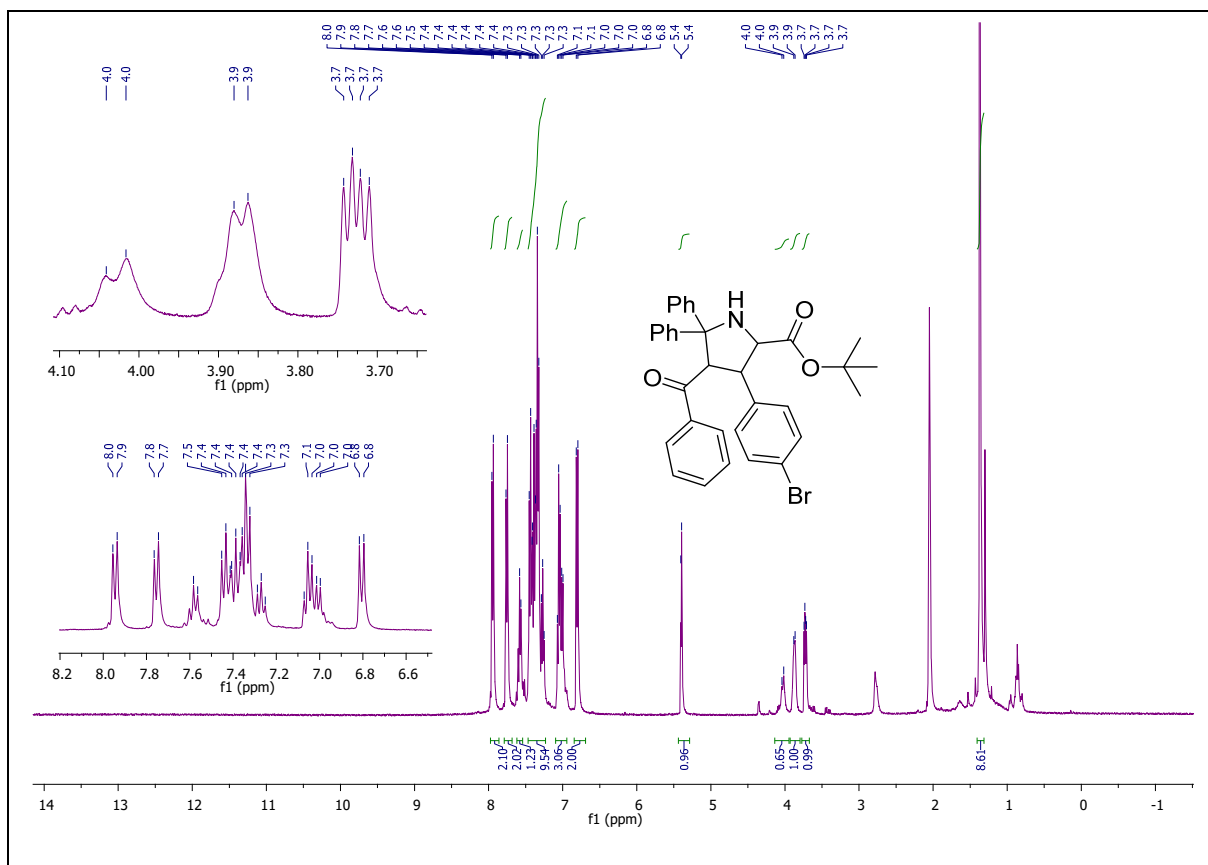


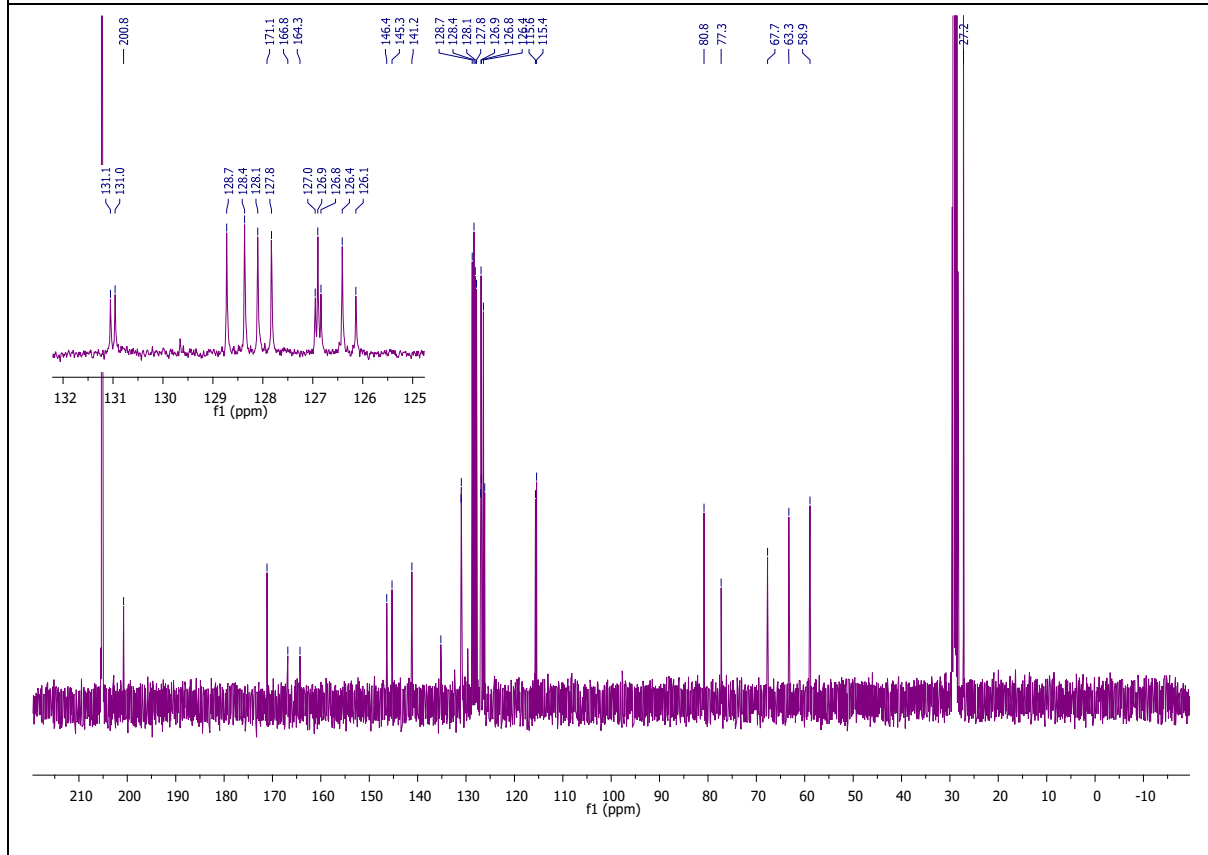
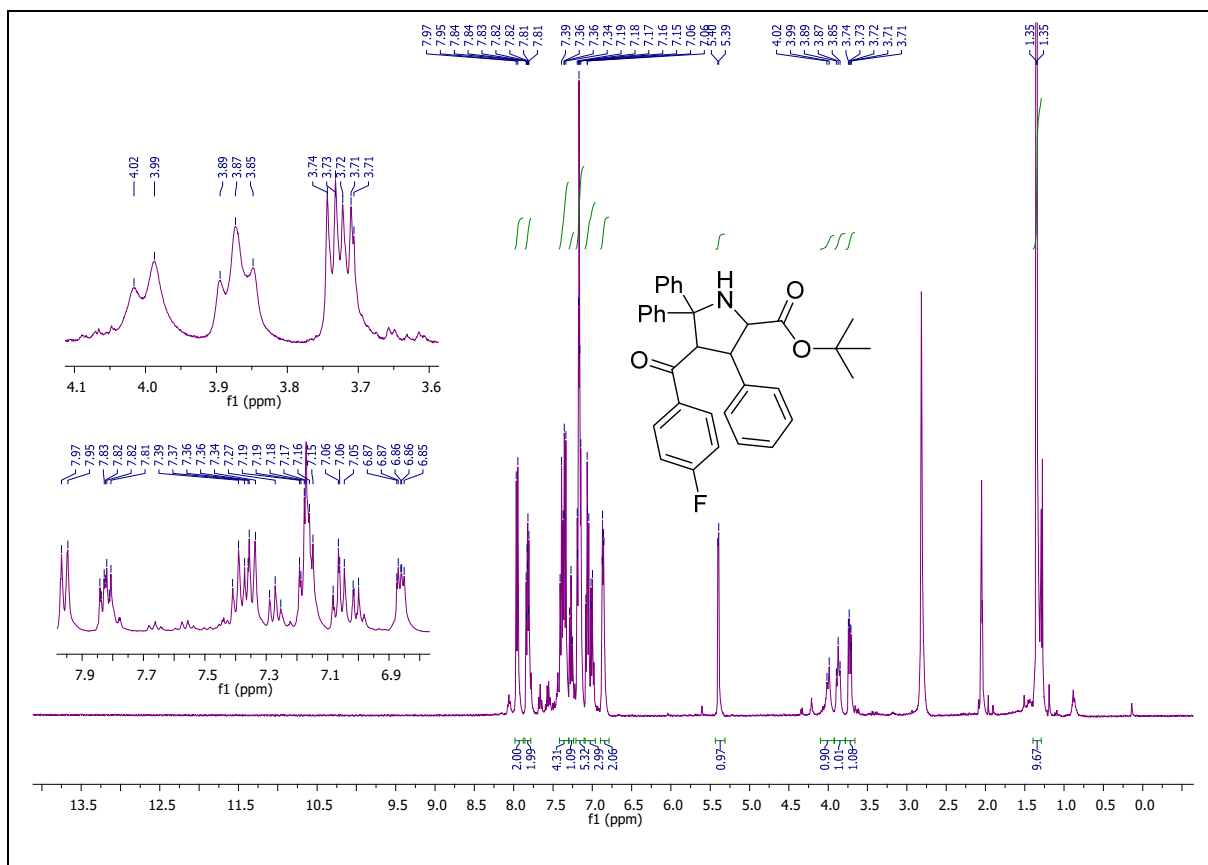


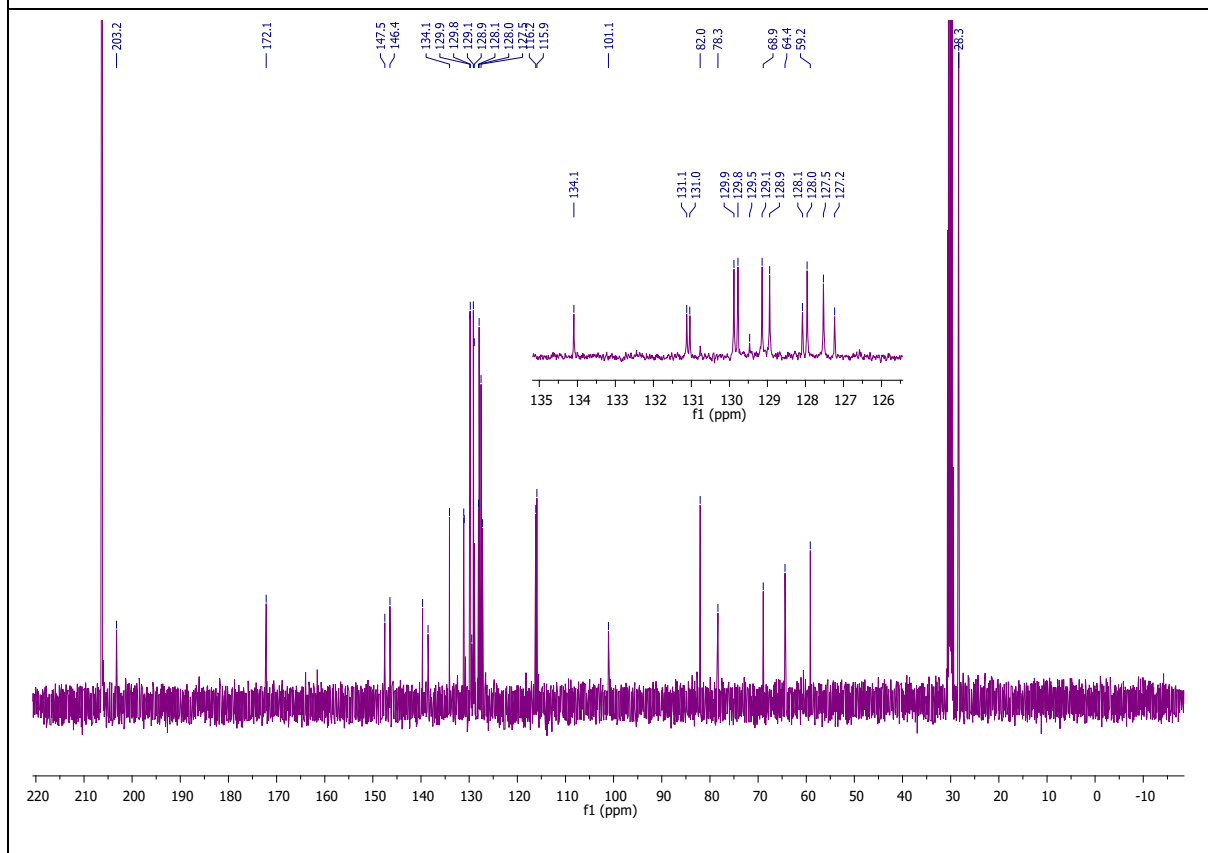
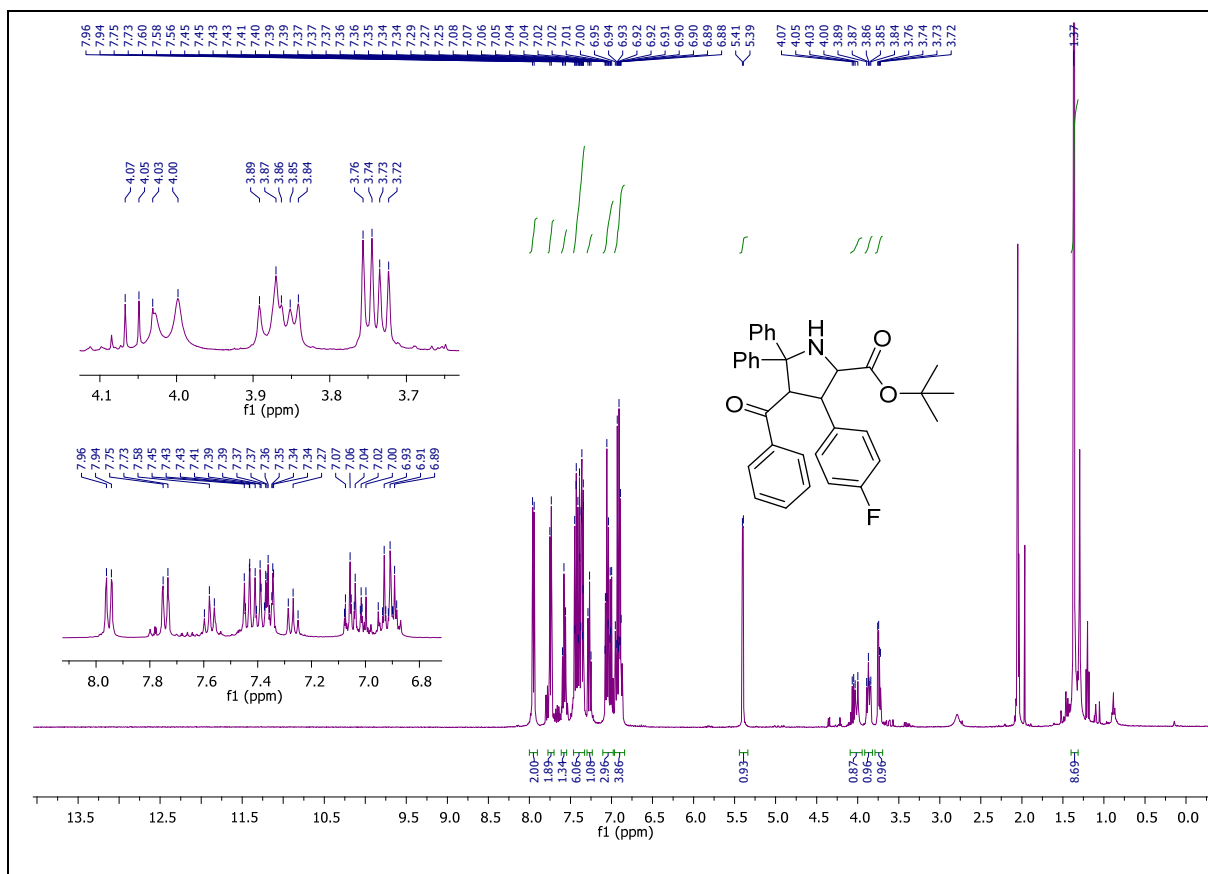


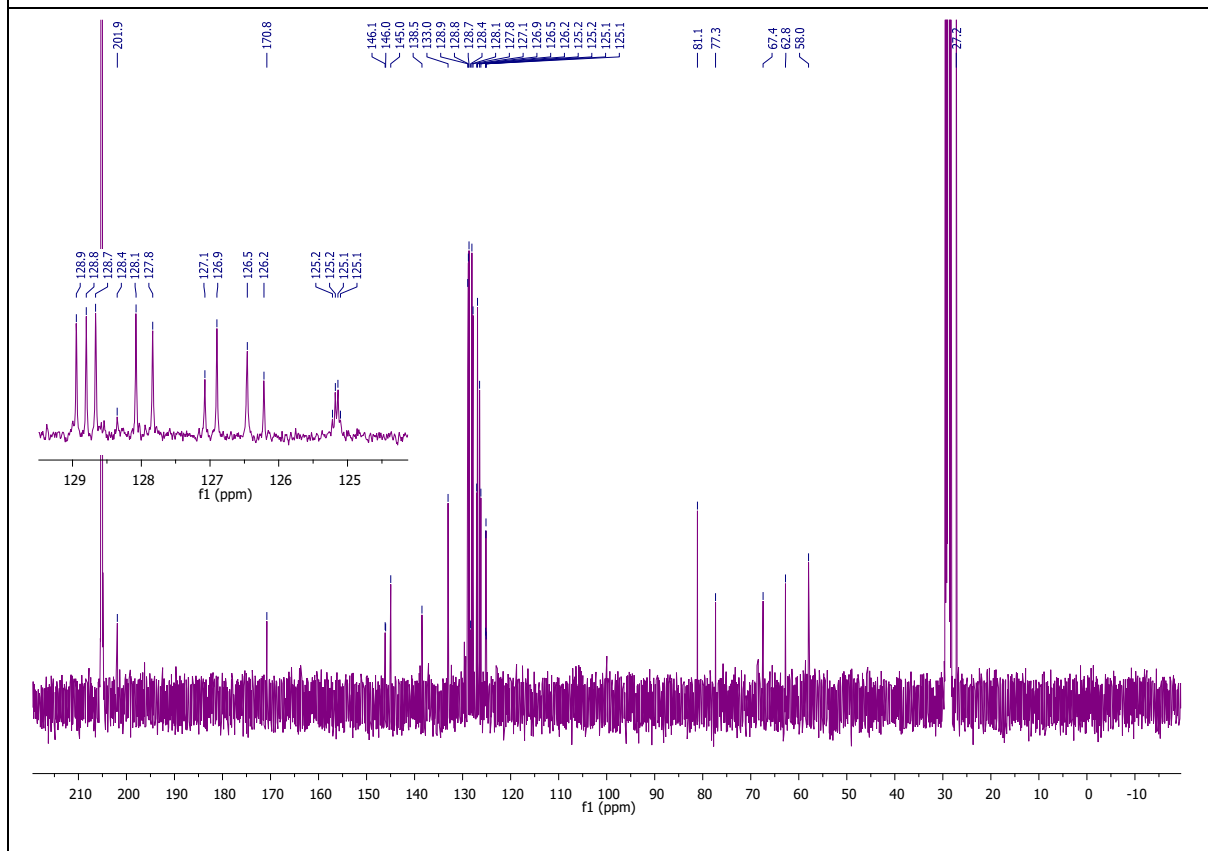
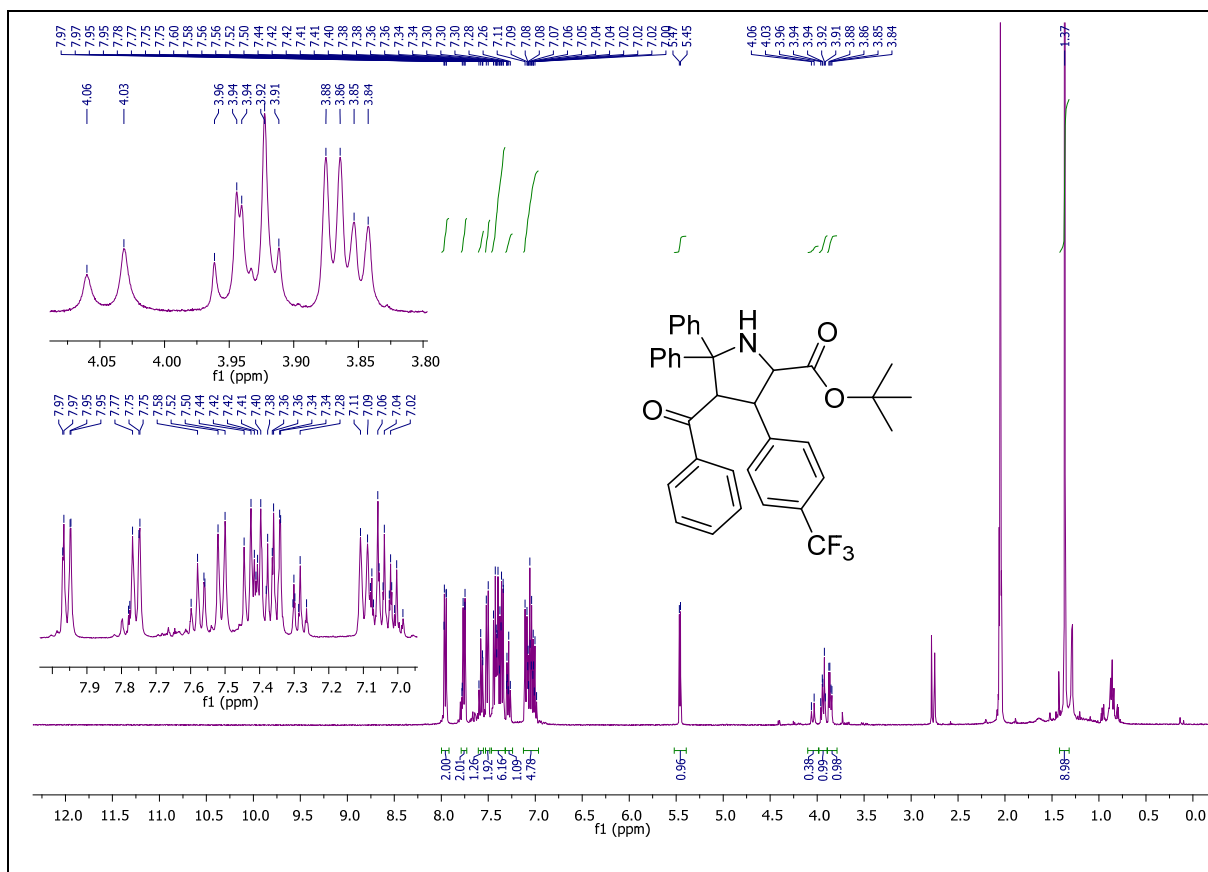


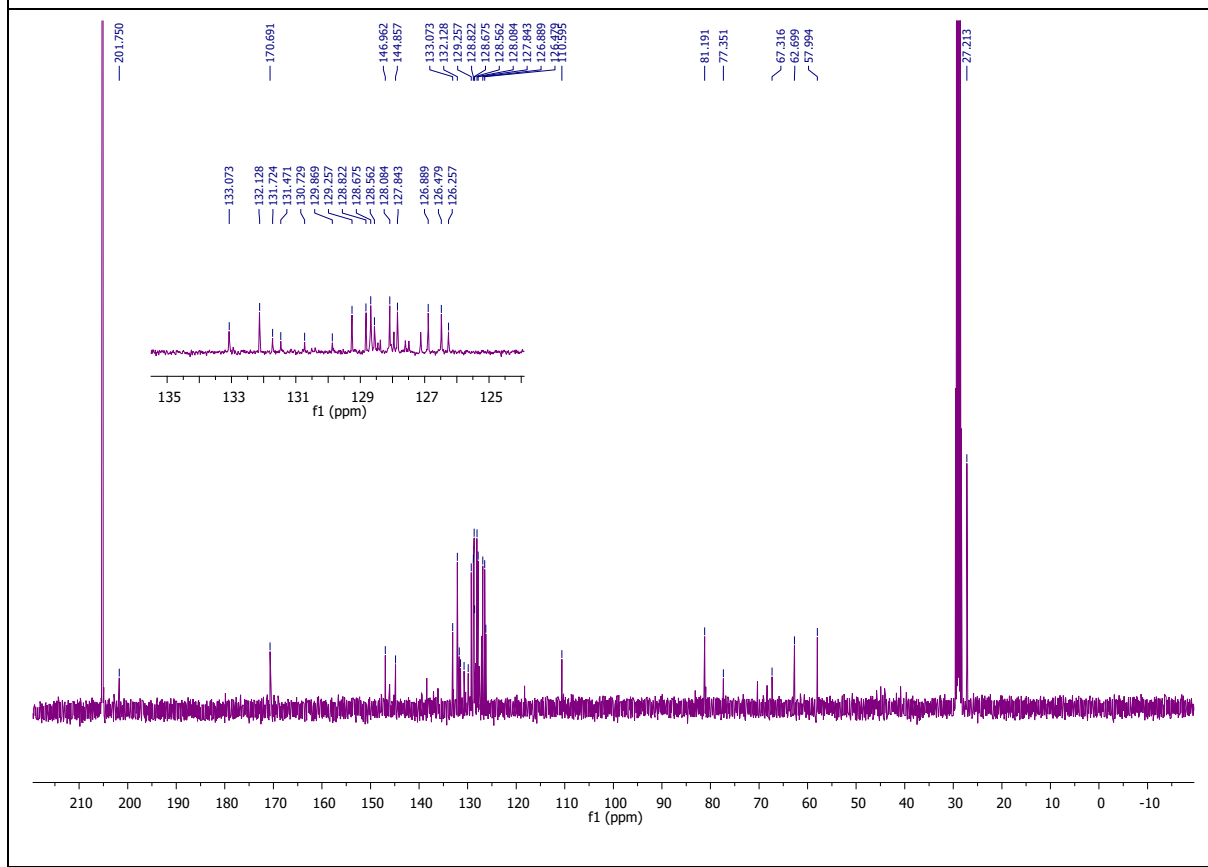
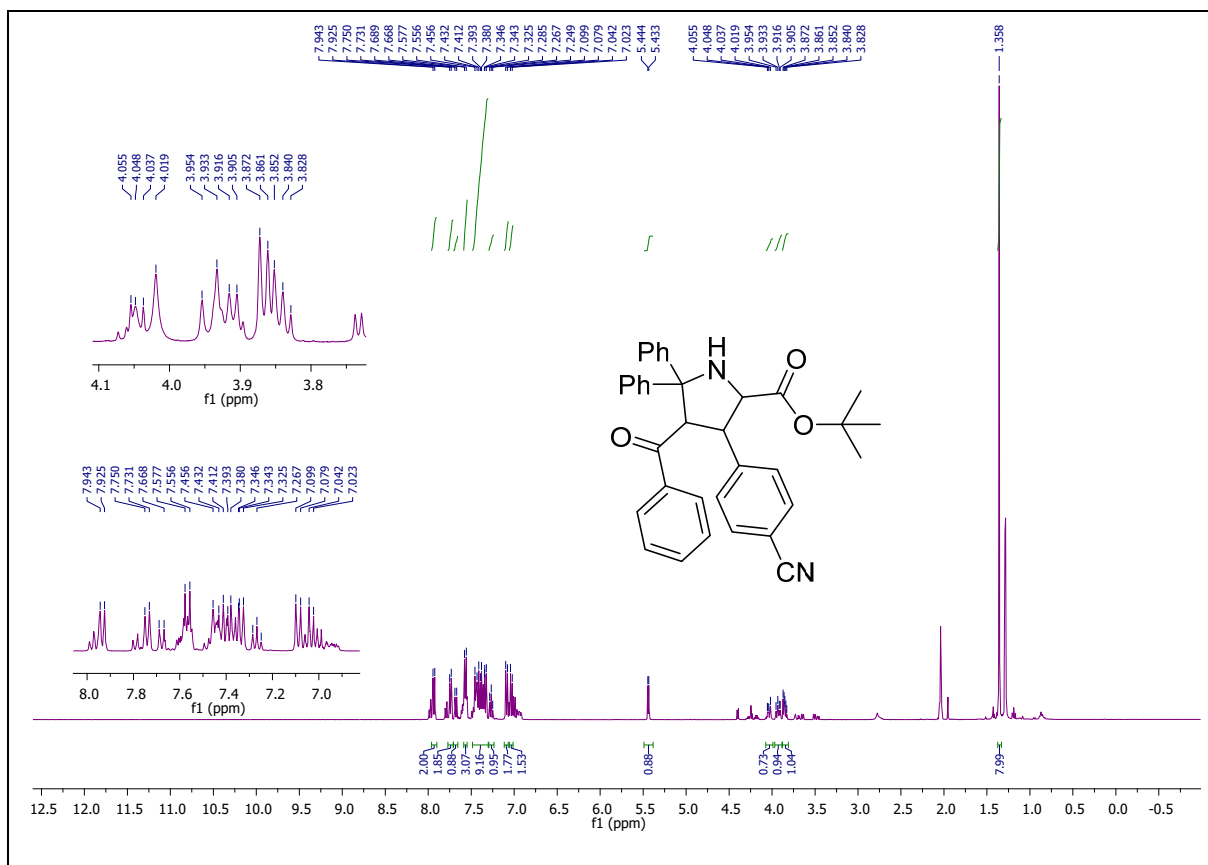


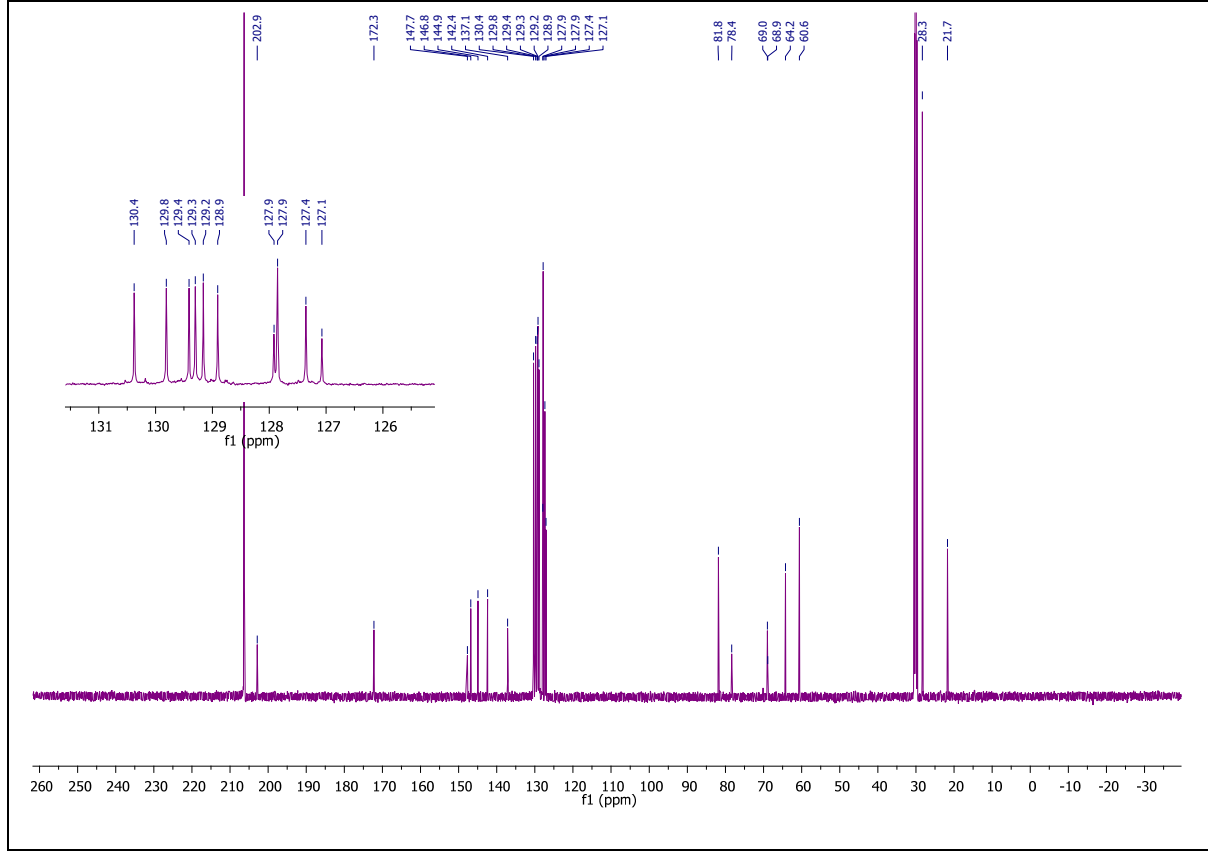
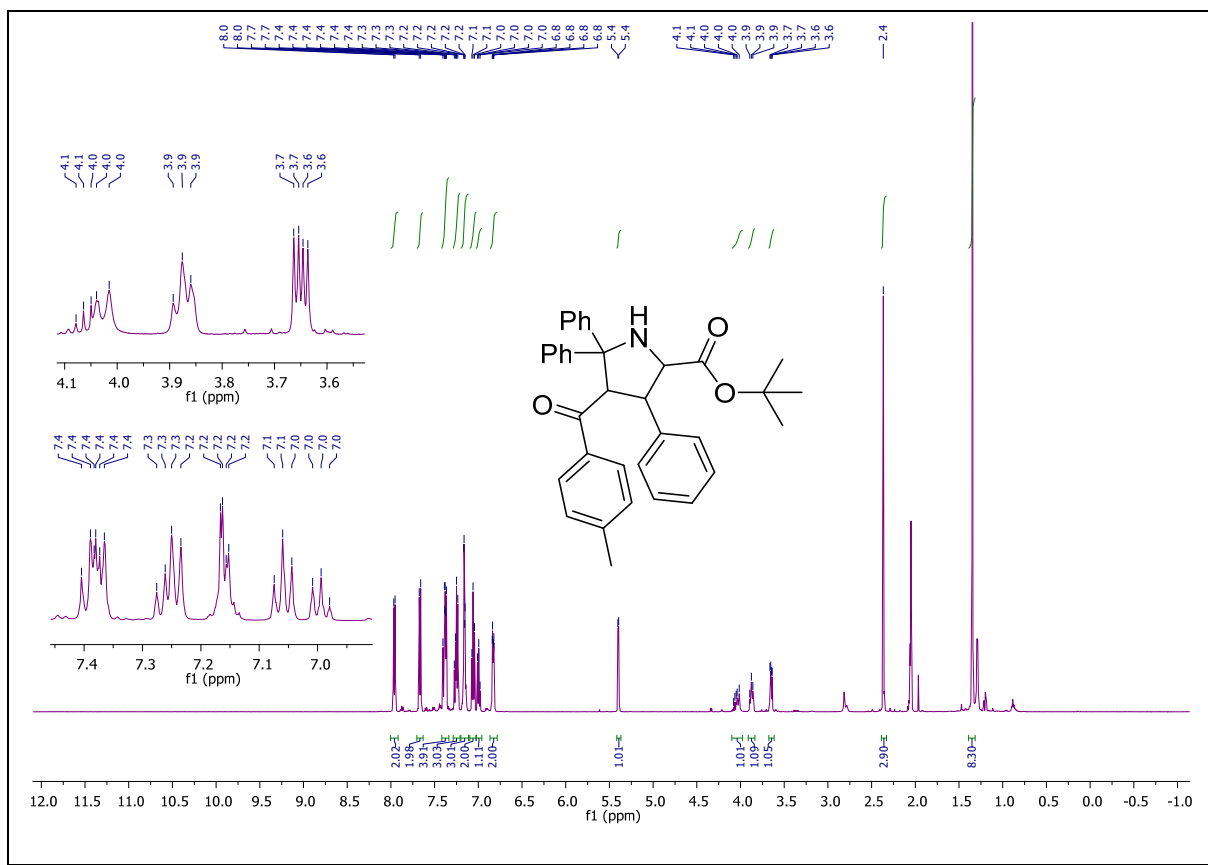


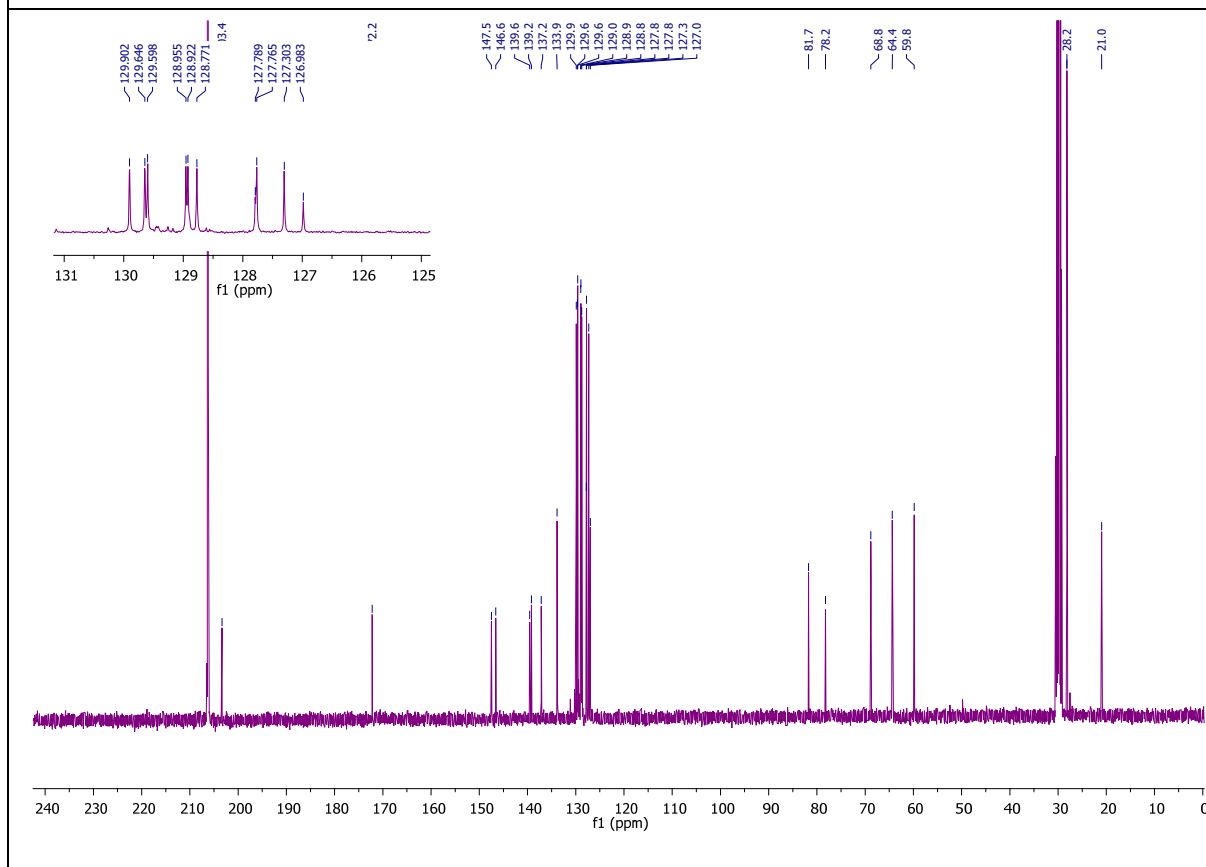
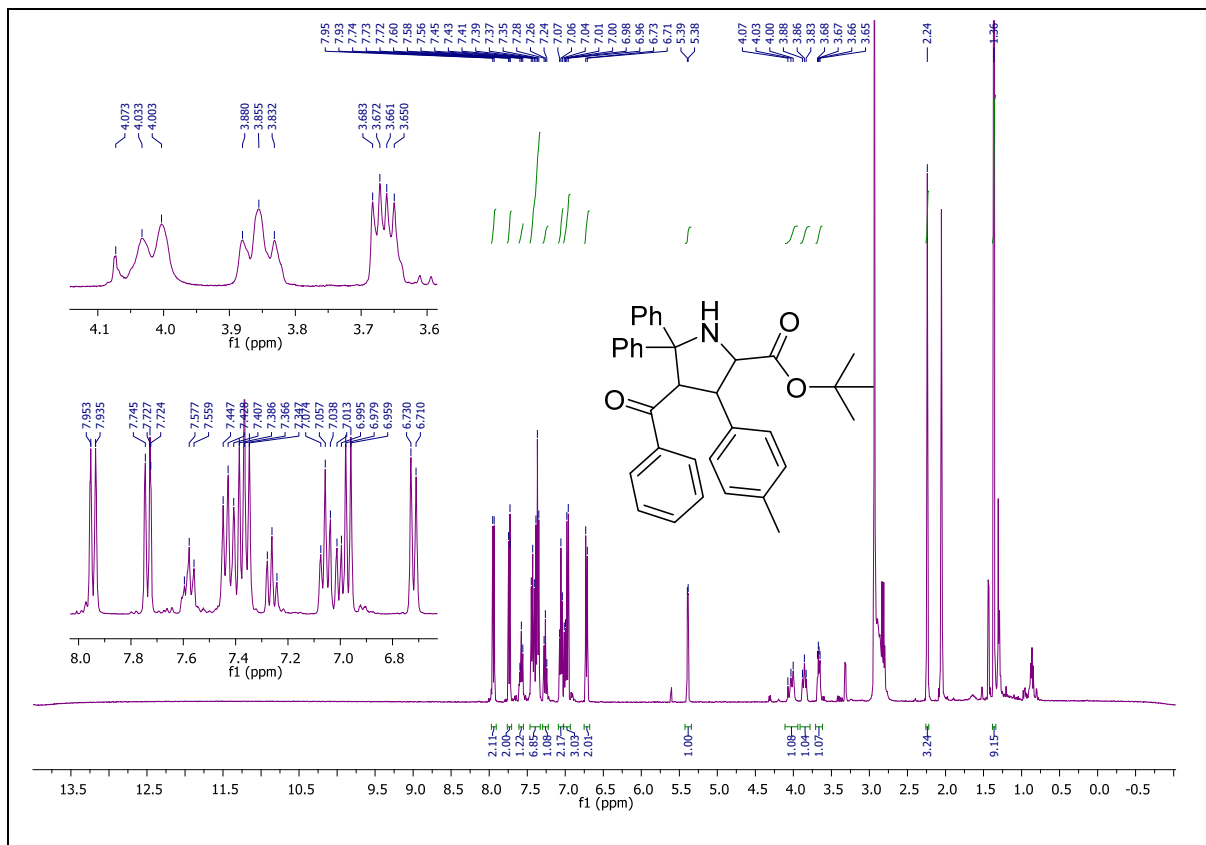


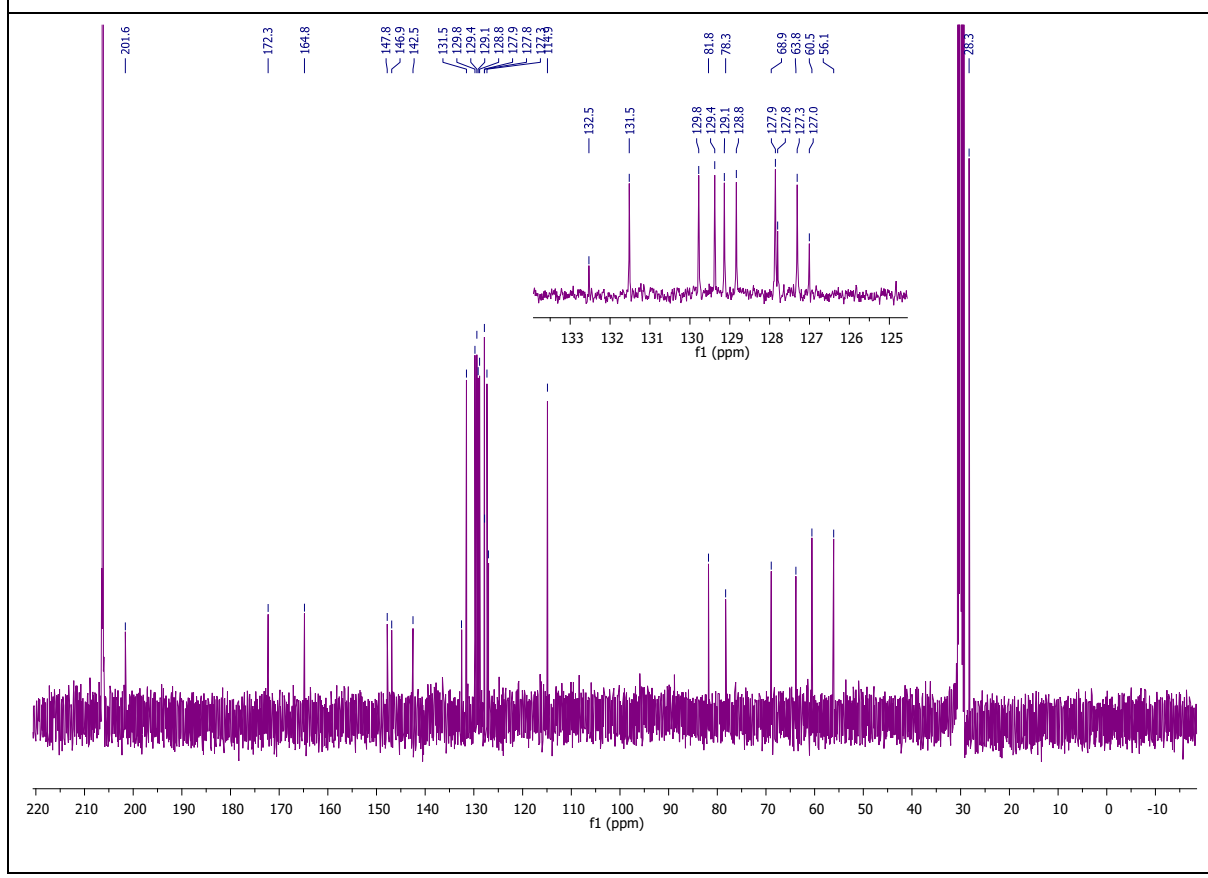
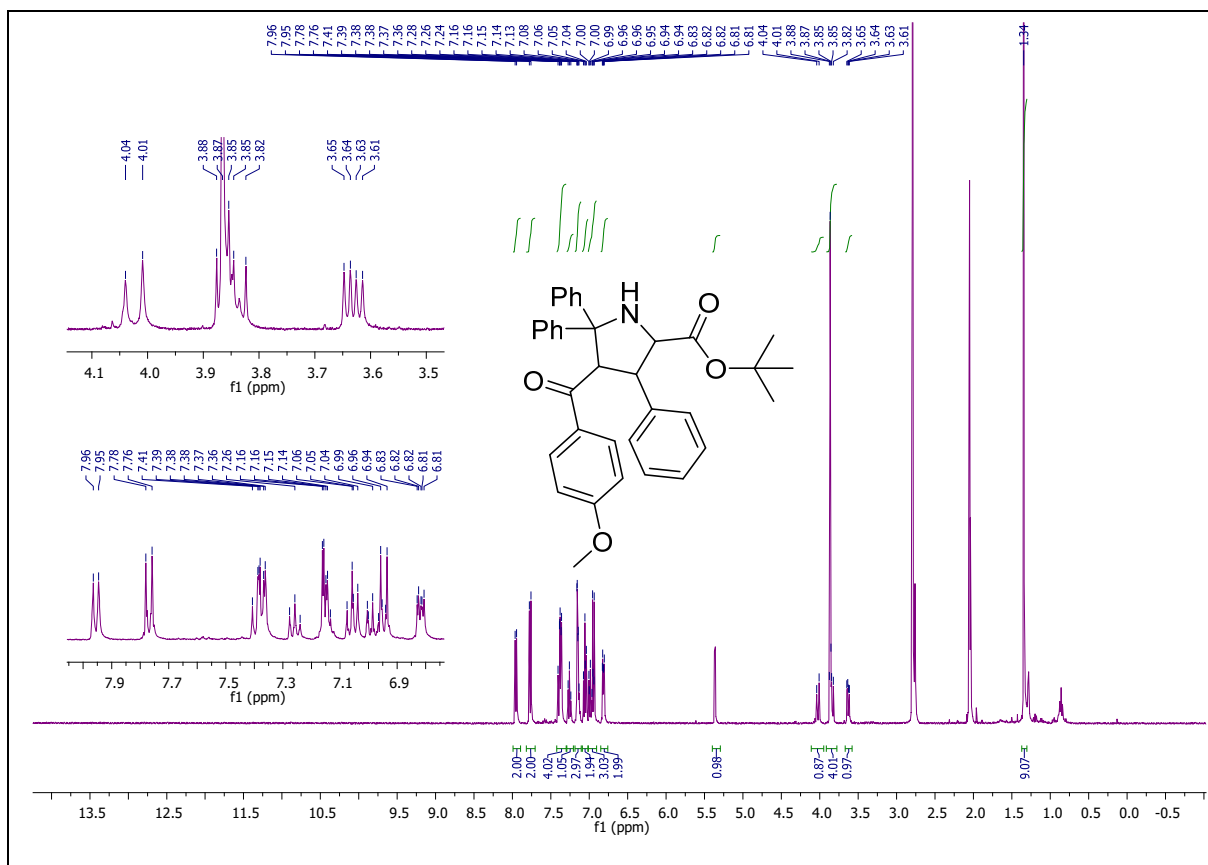


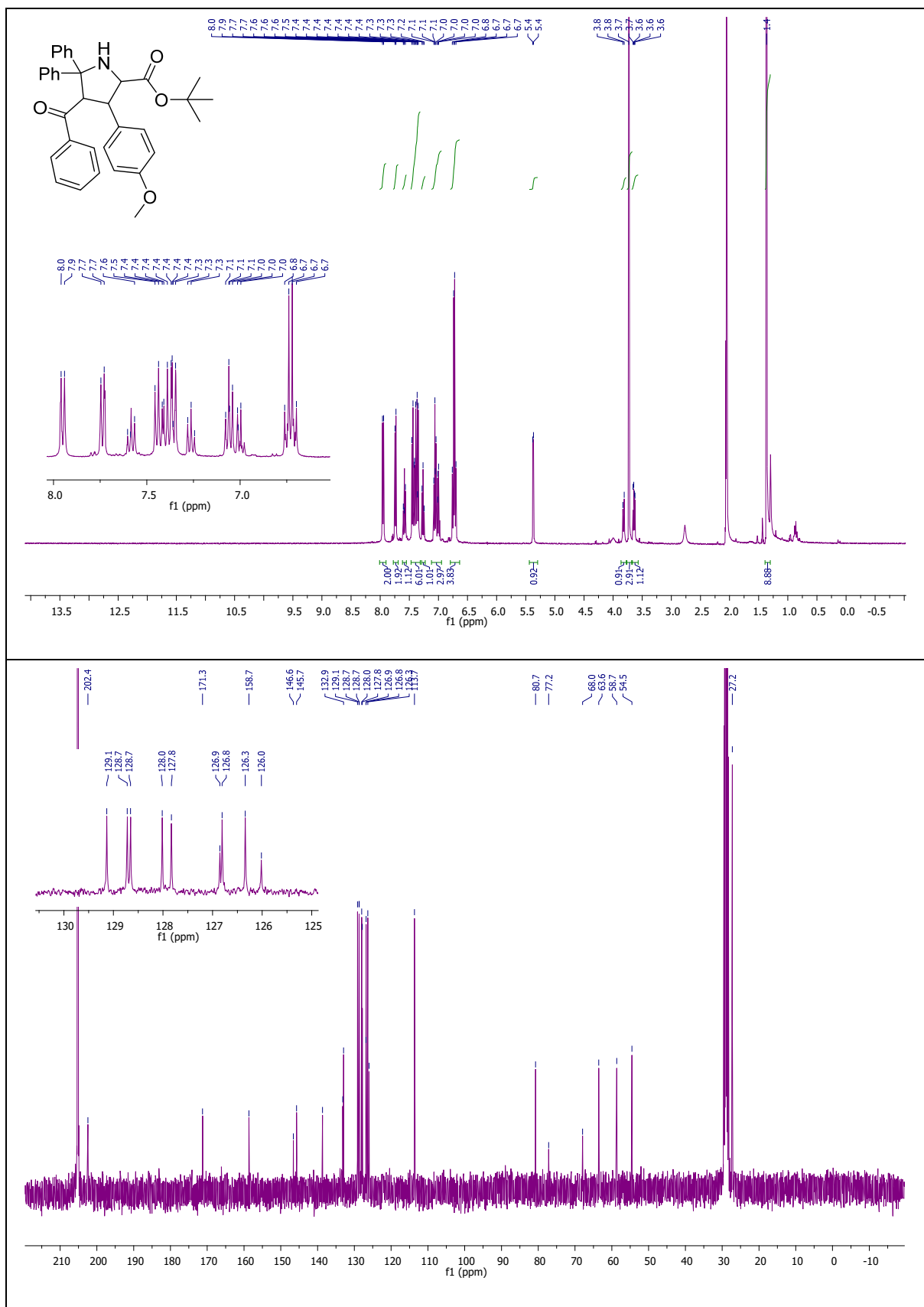


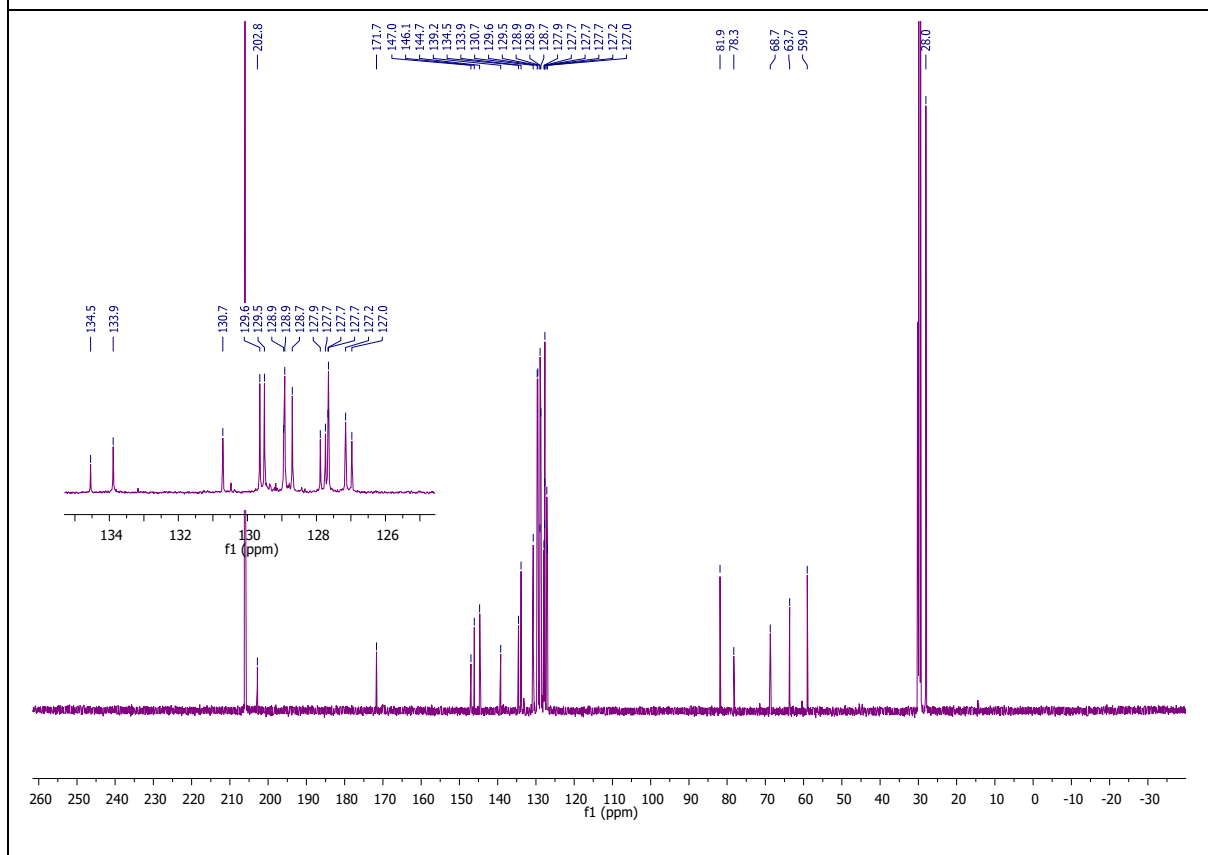
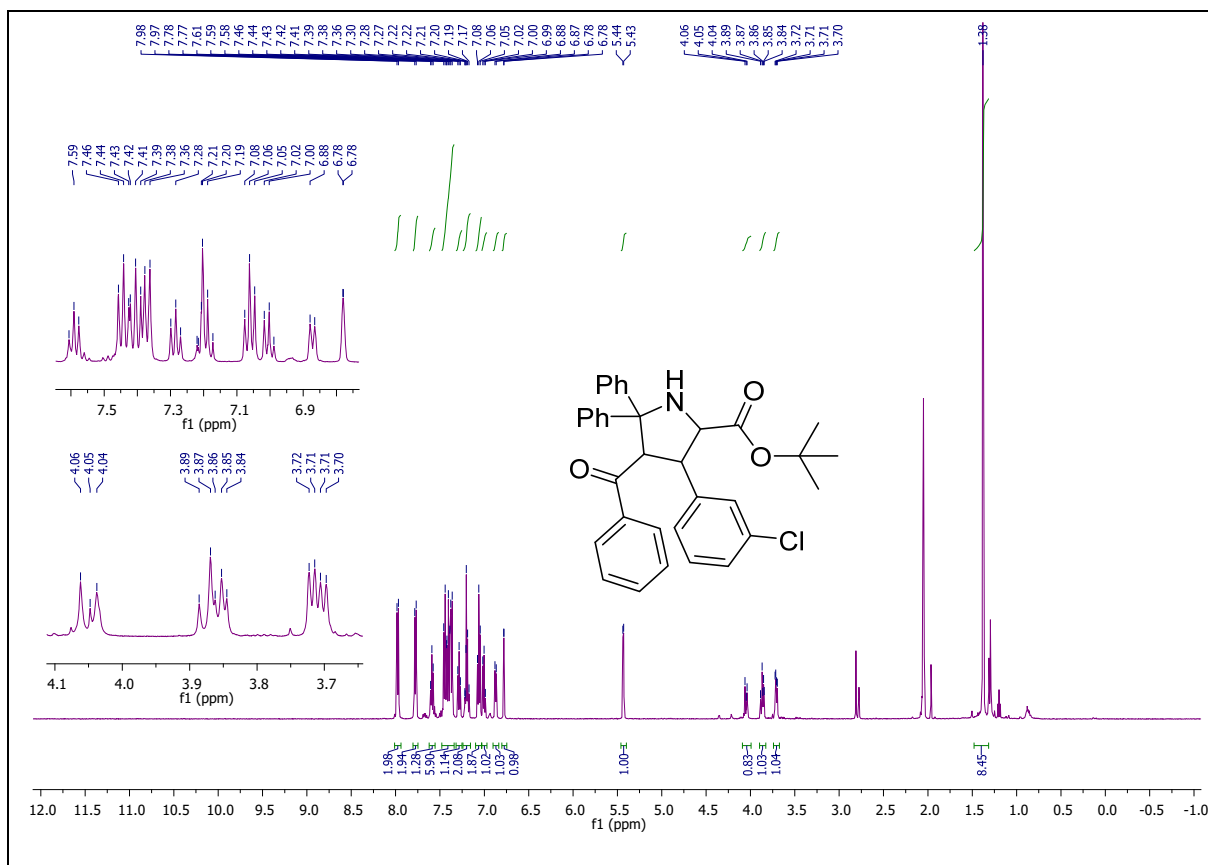


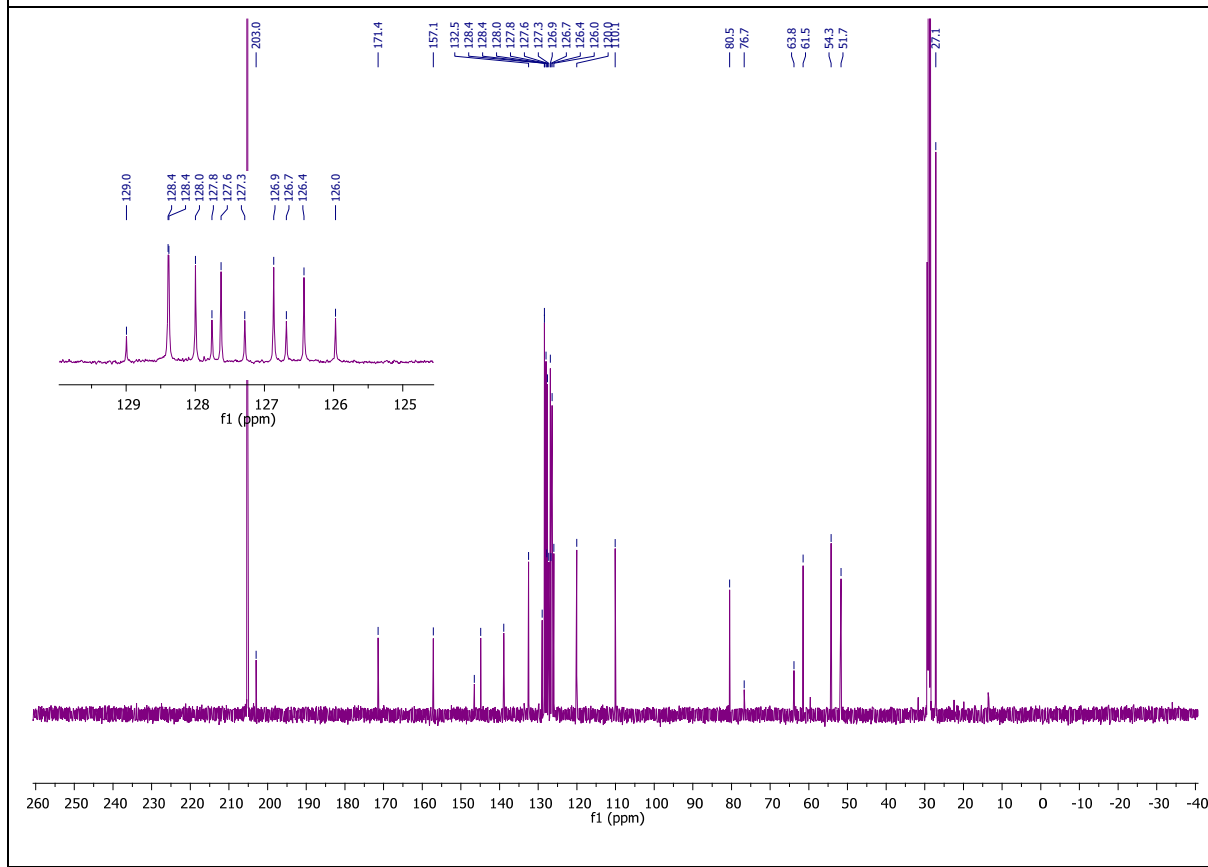
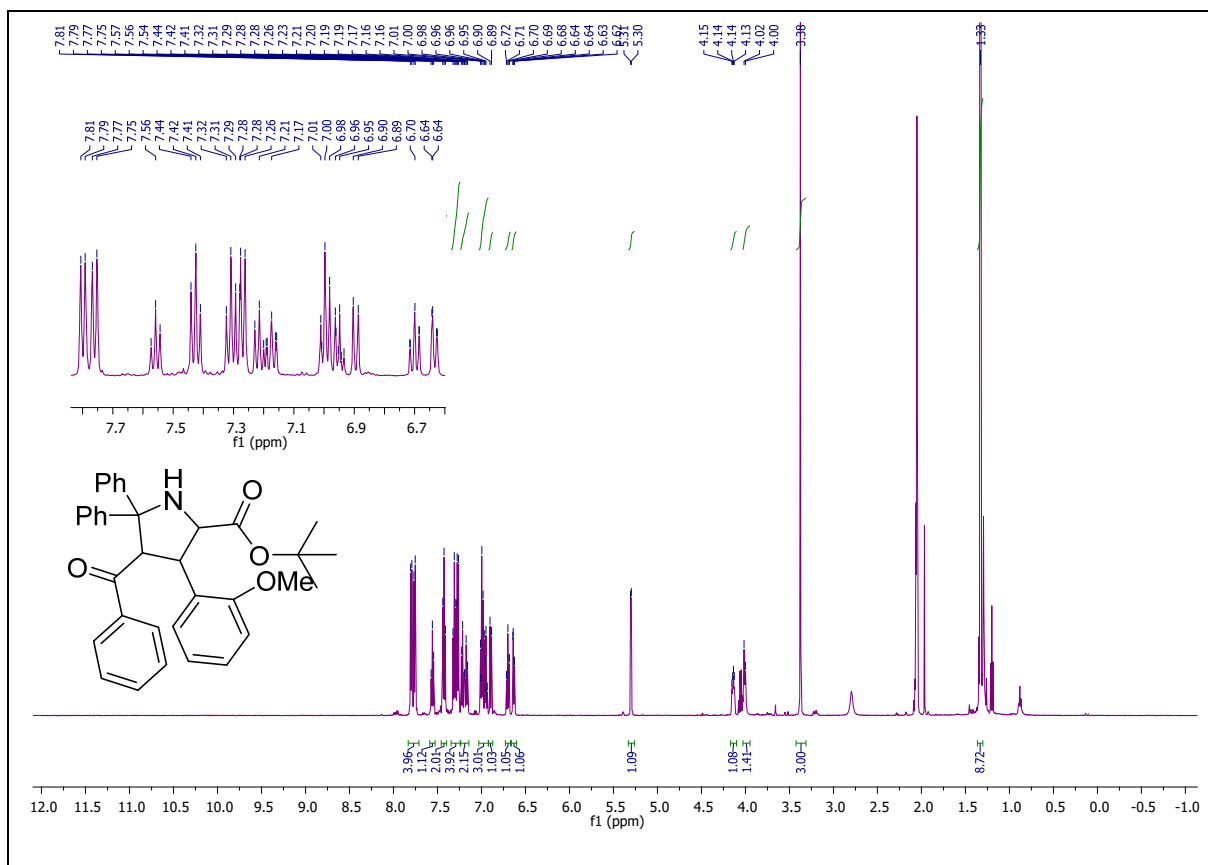


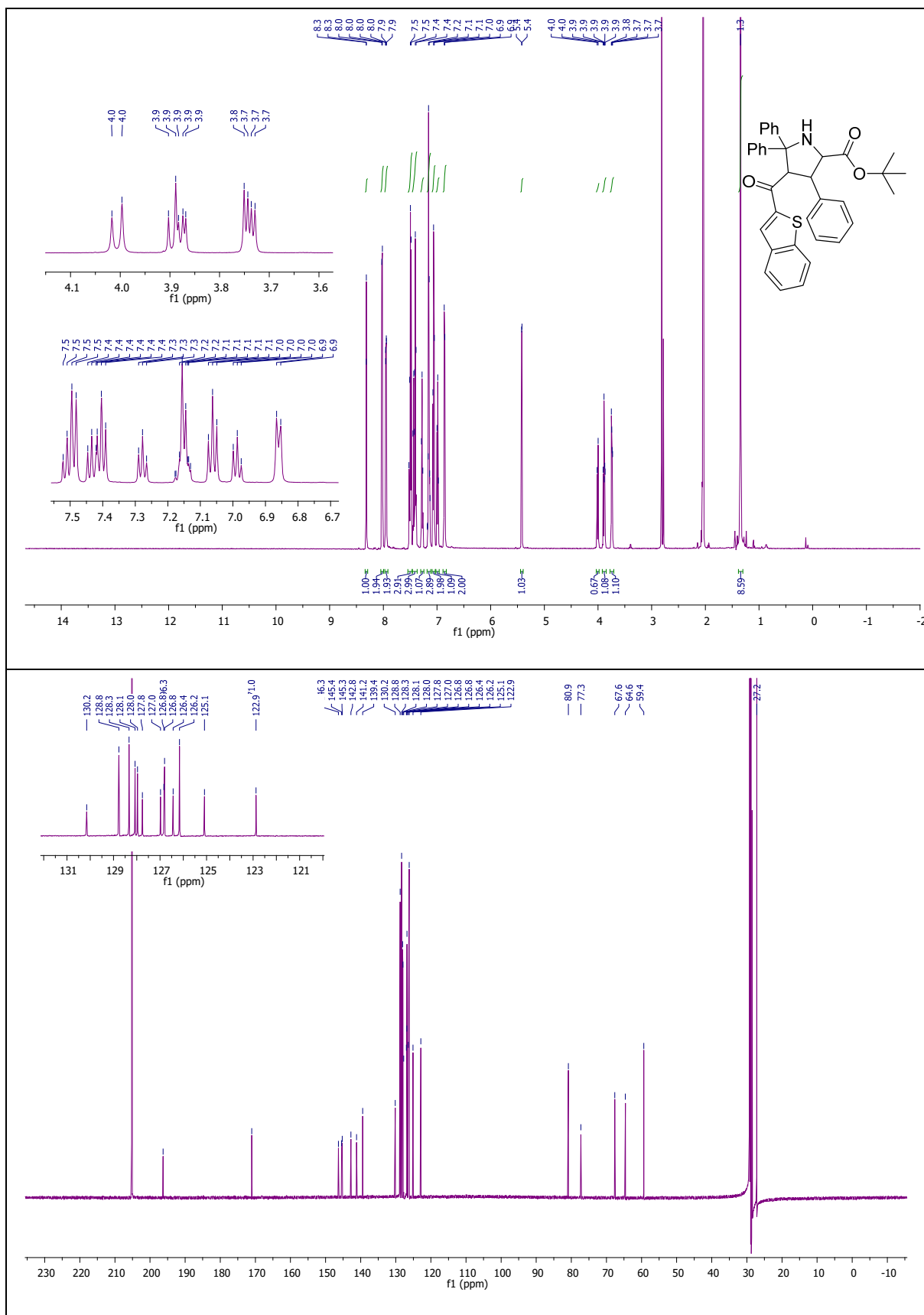


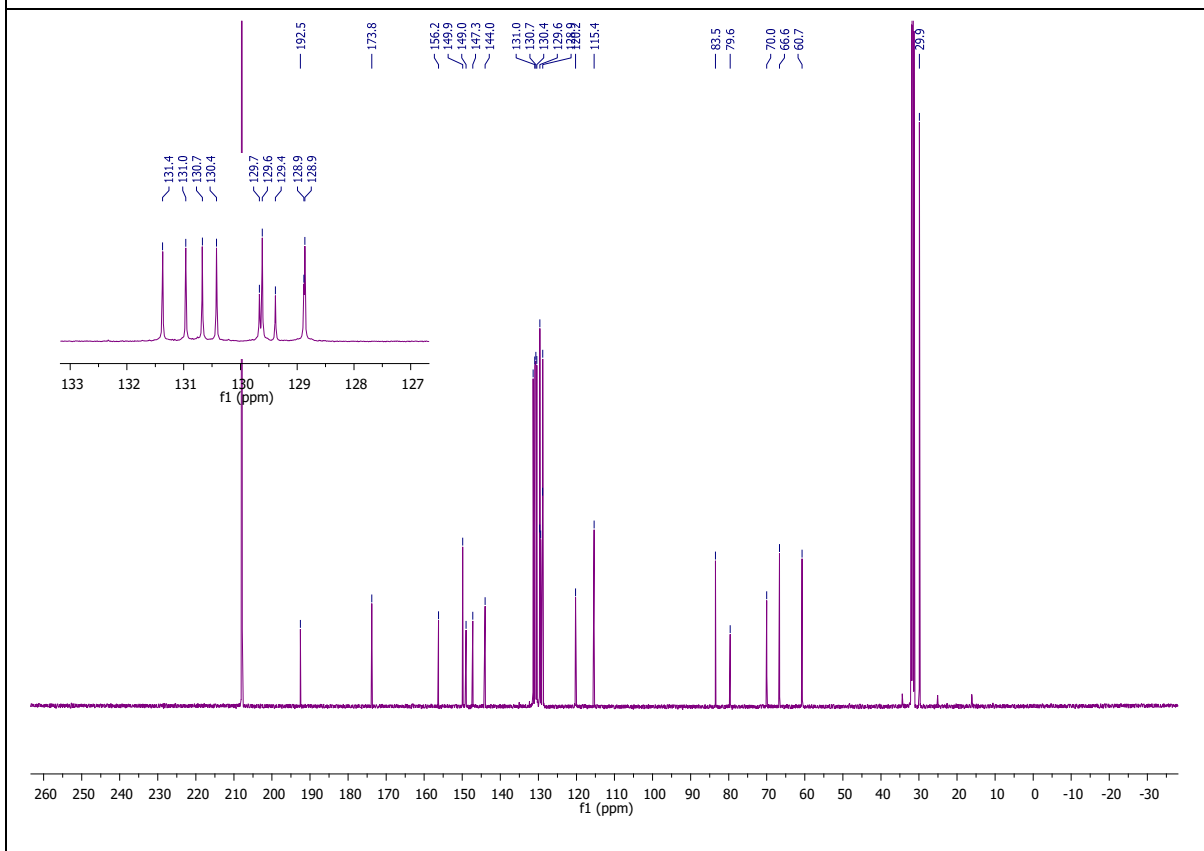
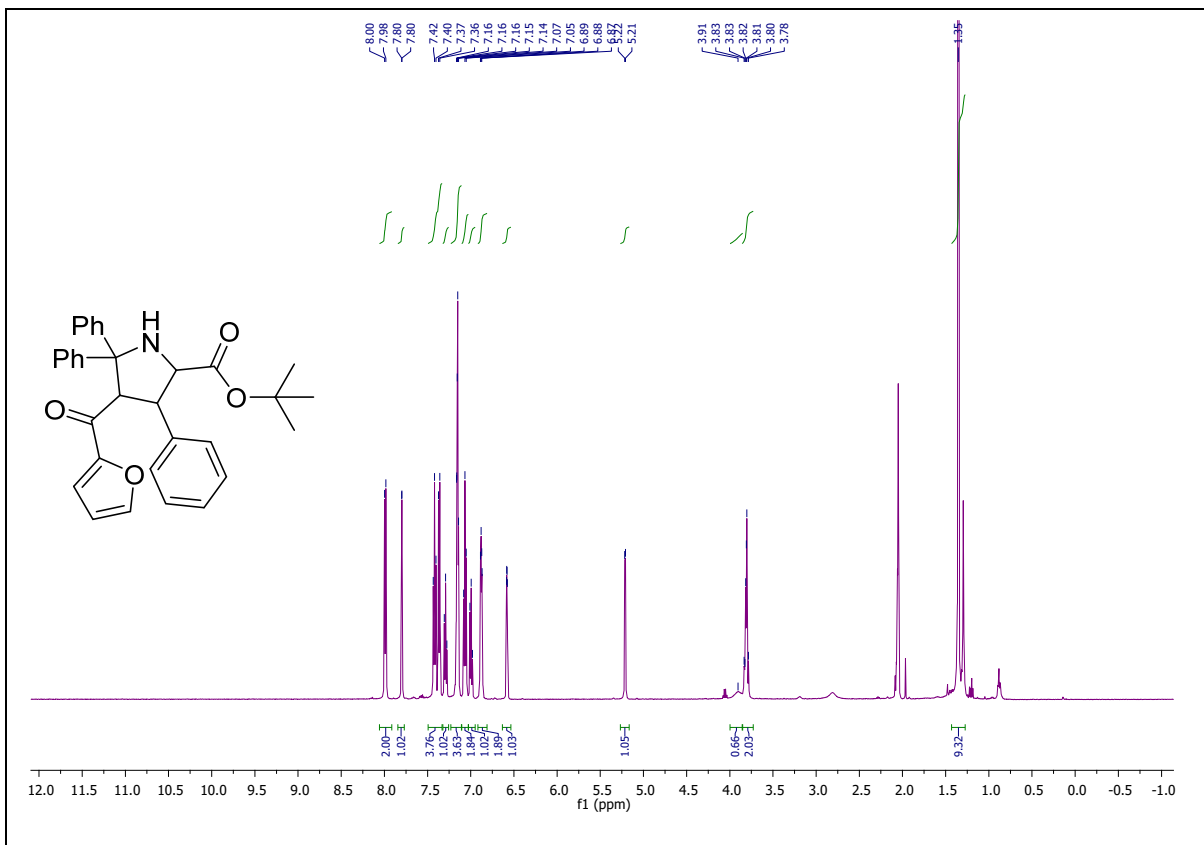


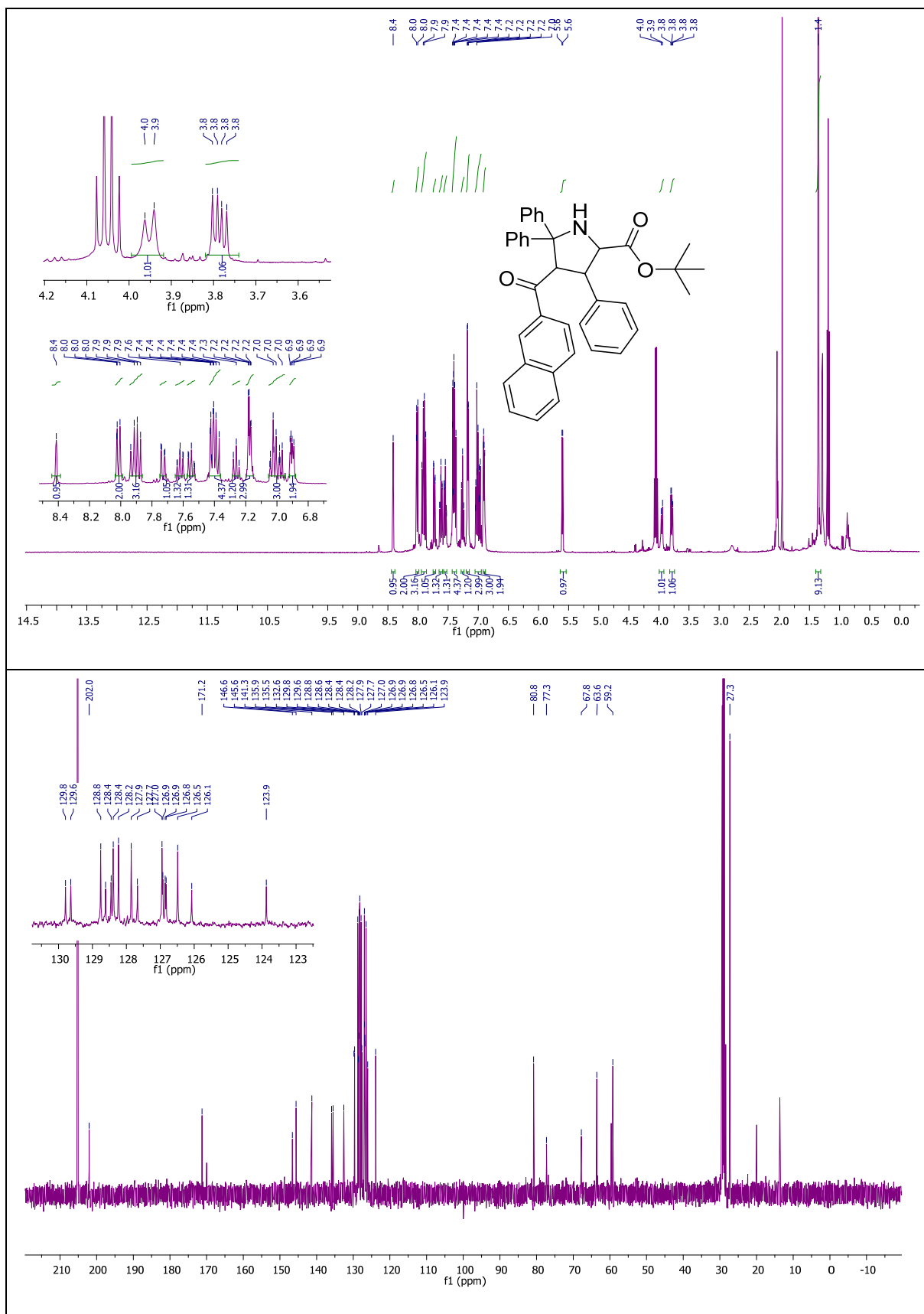


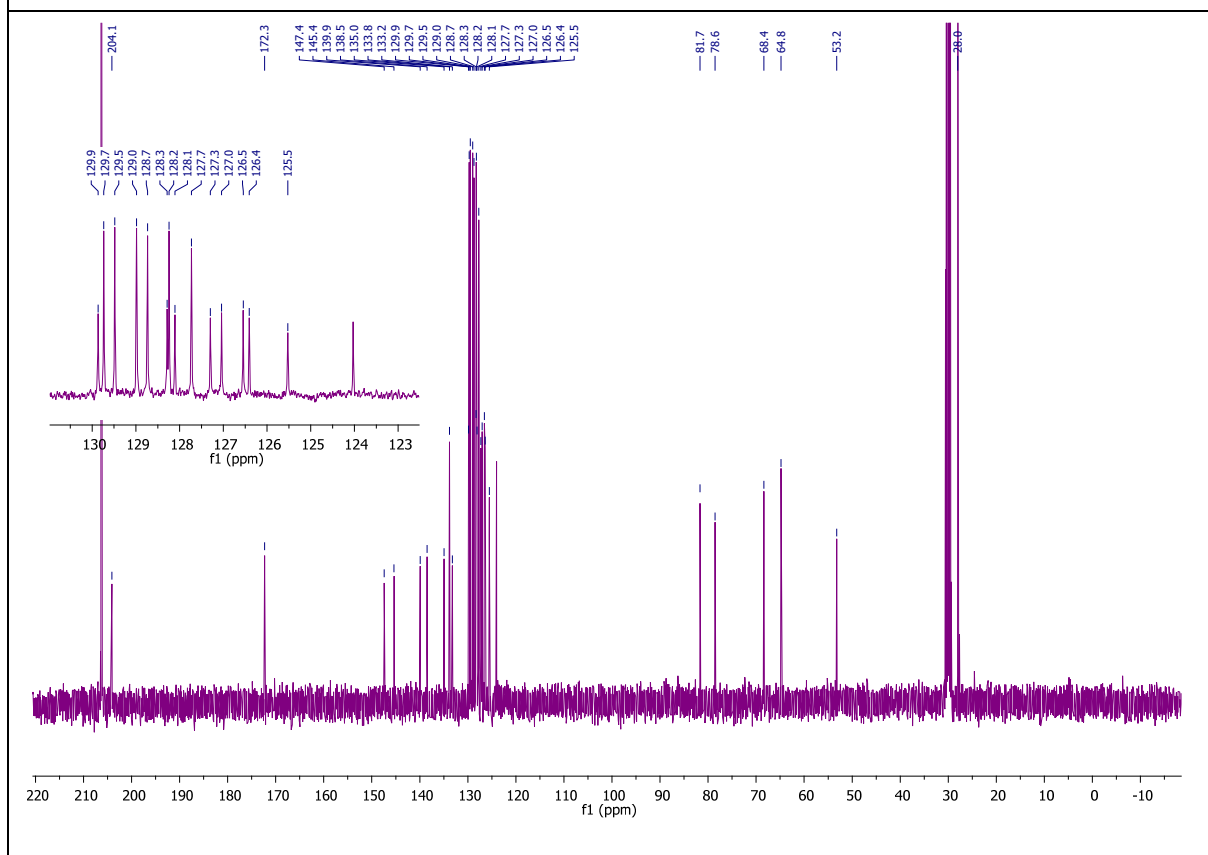
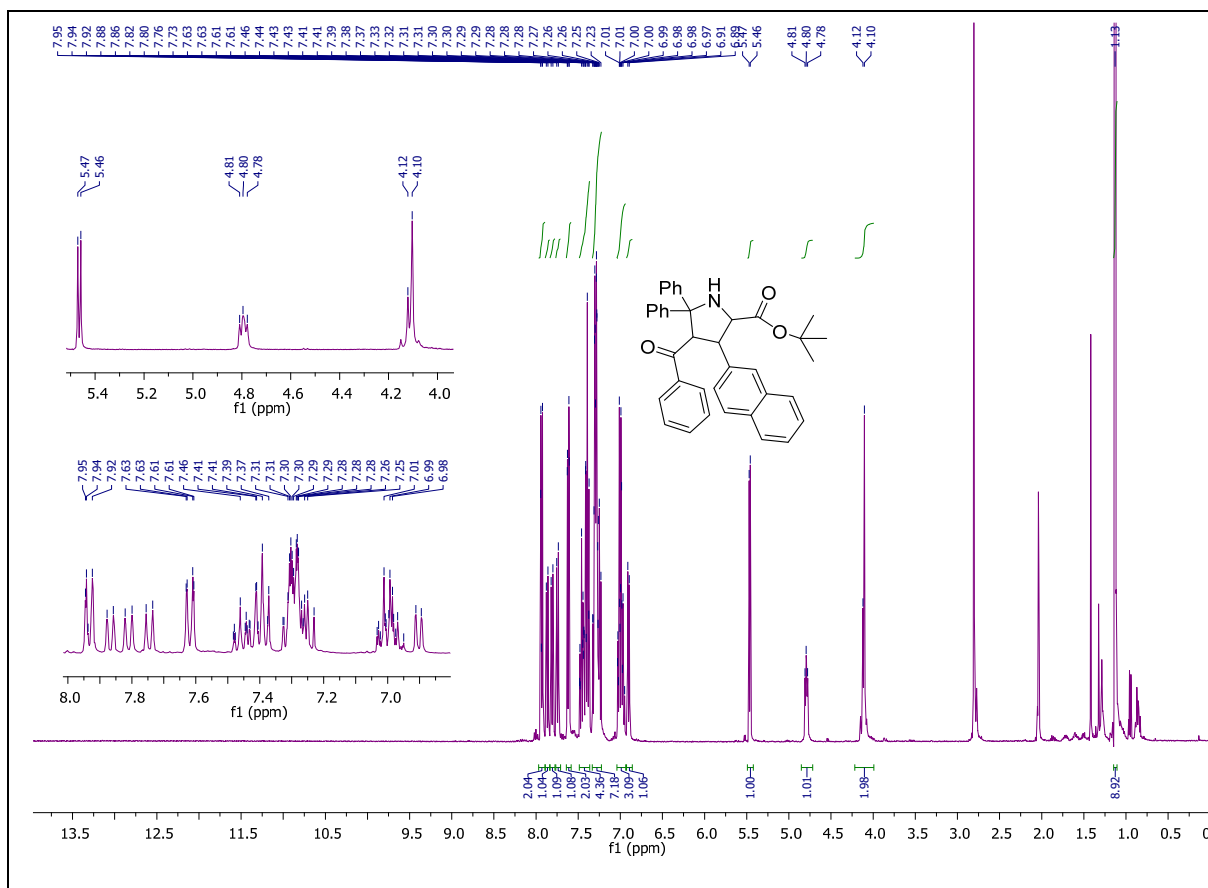




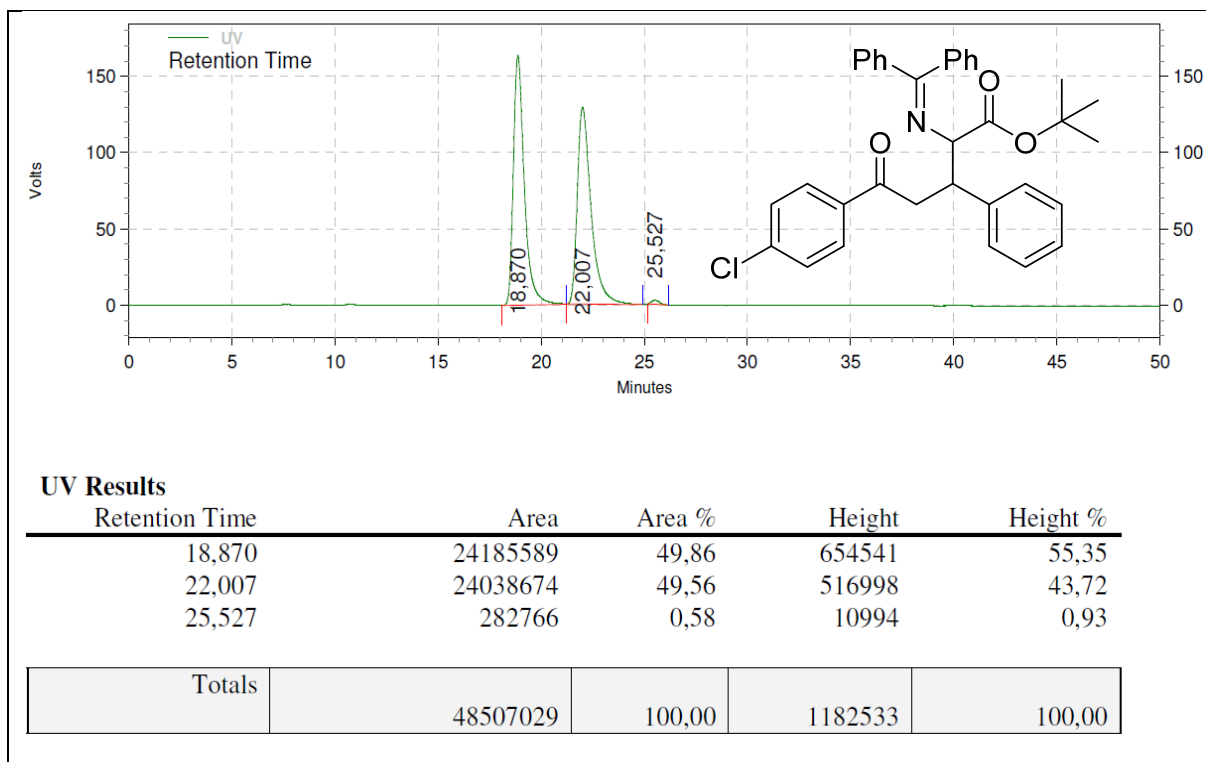
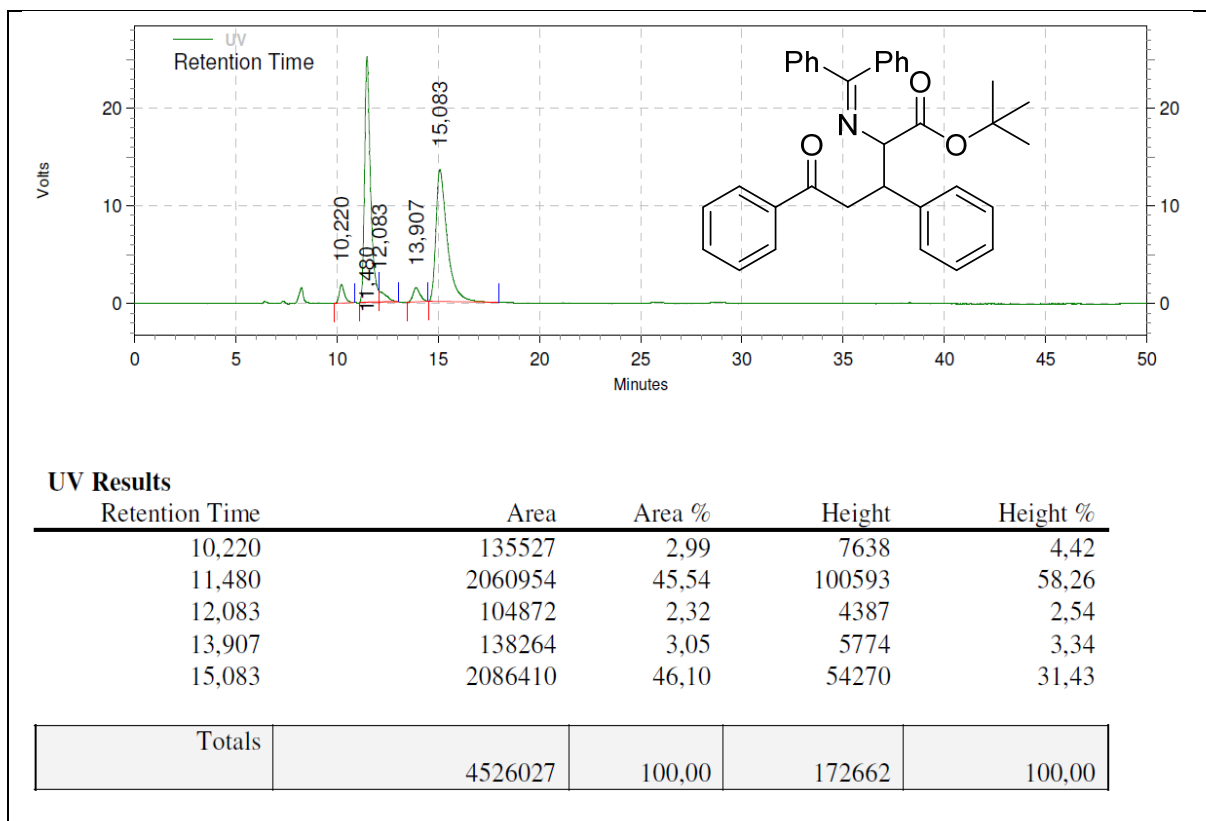


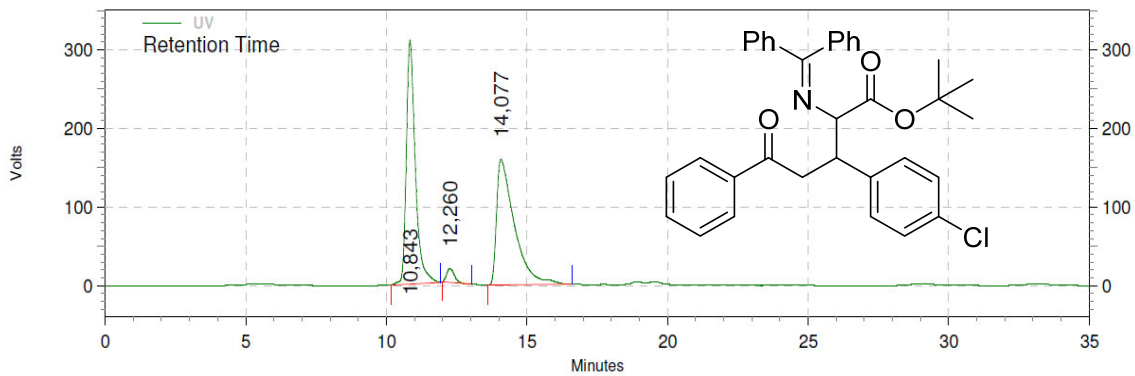






4. Copies of HPLC chromatograms

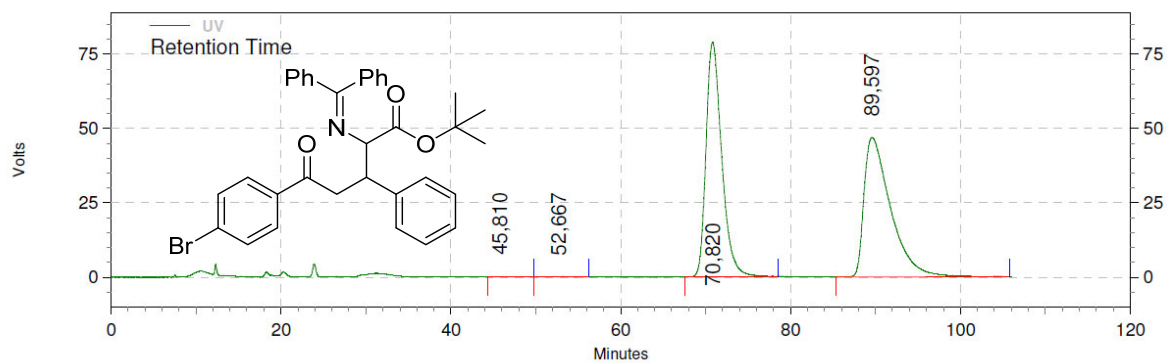




UV Results

Retention Time	Area	Area %	Height	Height %
10,843	27505736	48,28	1242523	63,59
12,260	1340855	2,35	70533	3,61
14,077	28128019	49,37	640982	32,80

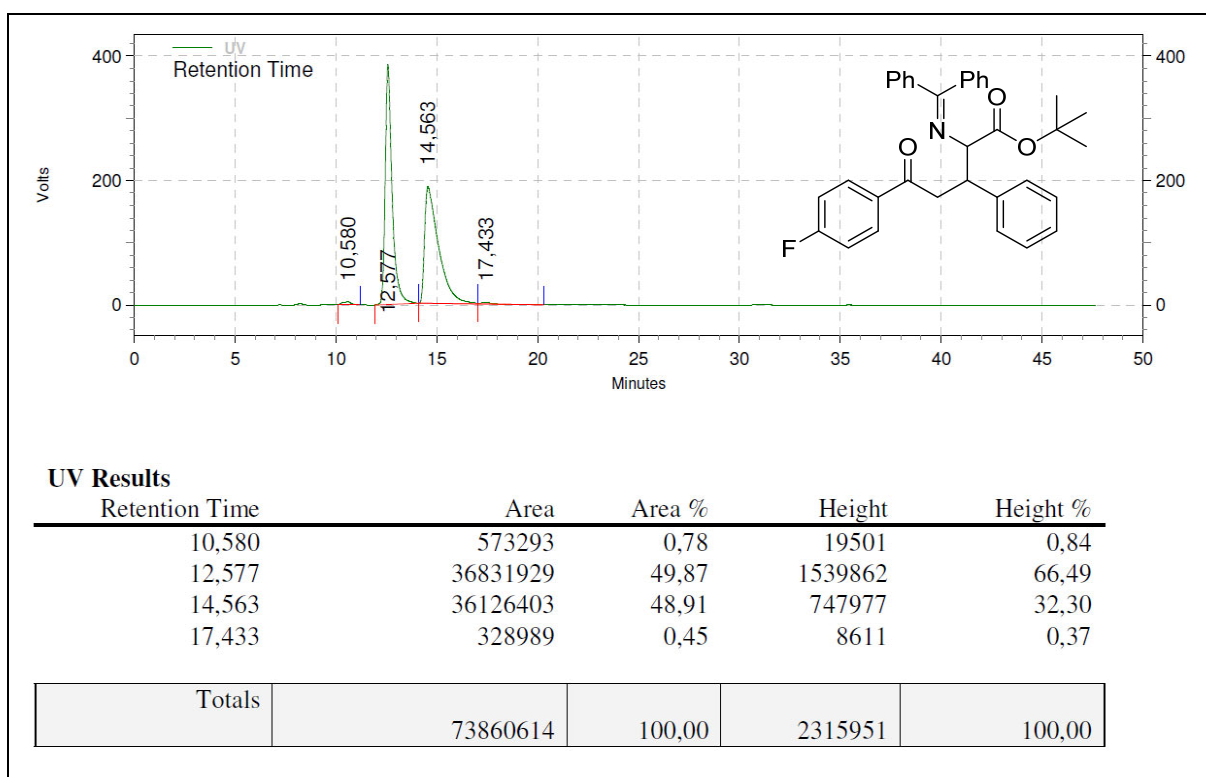
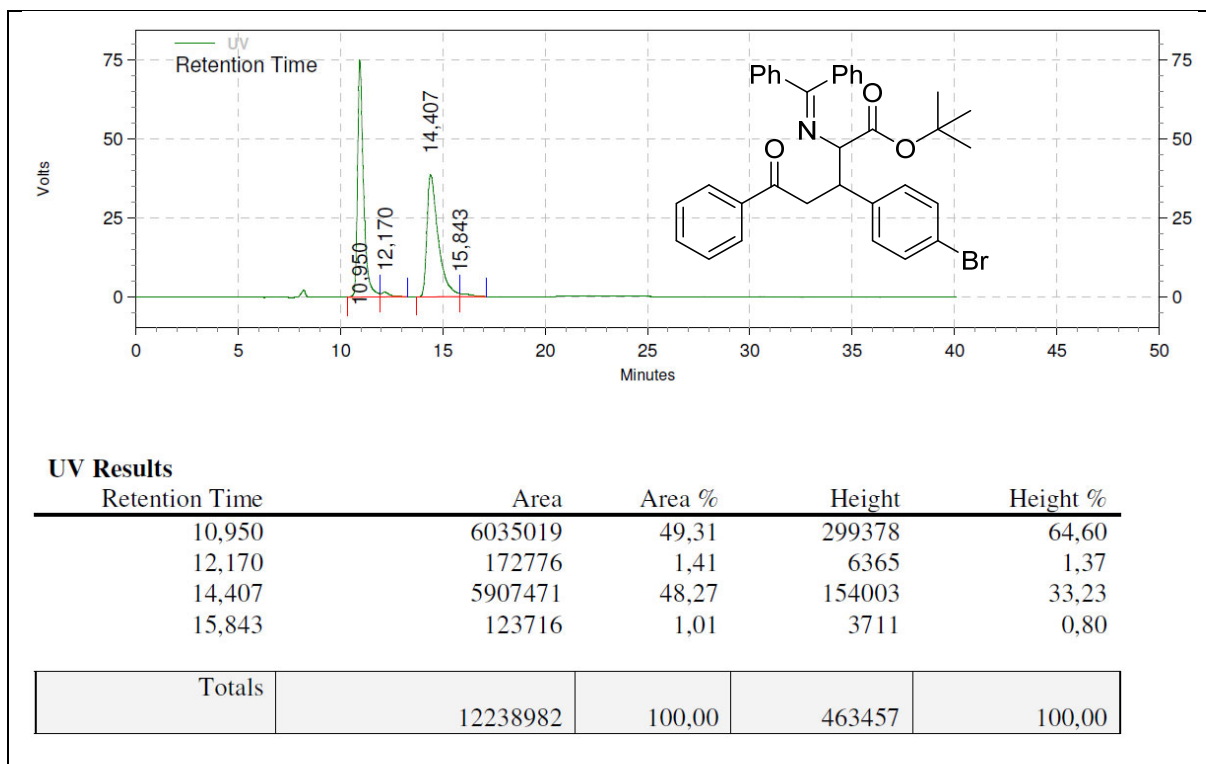
Totals	56974610	100,00	1954038	100,00
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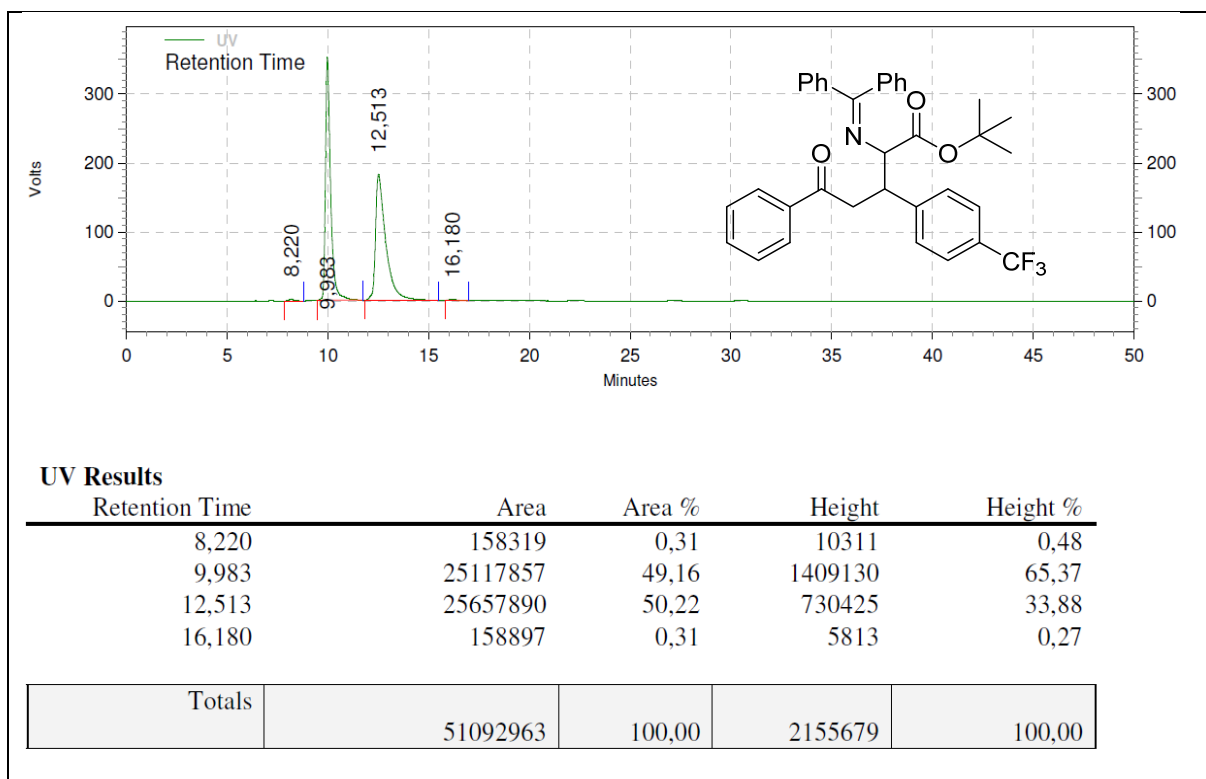
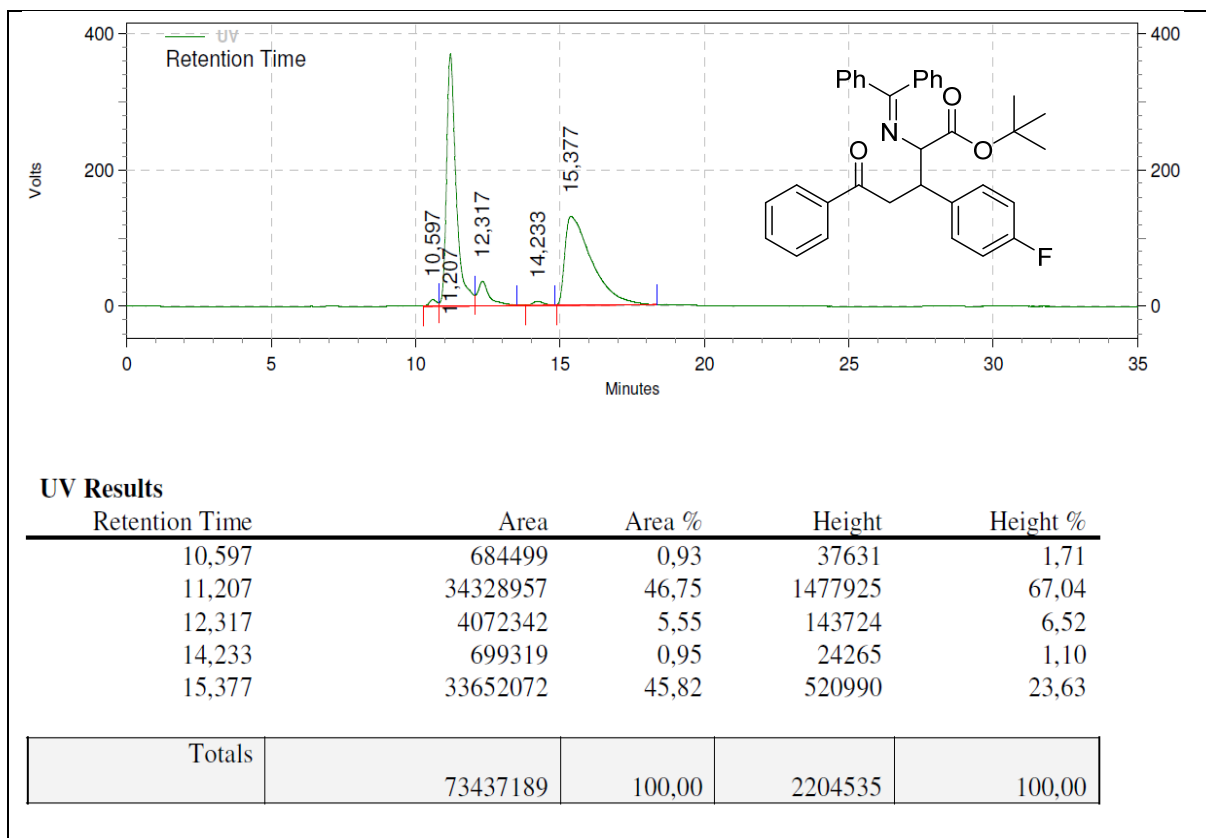


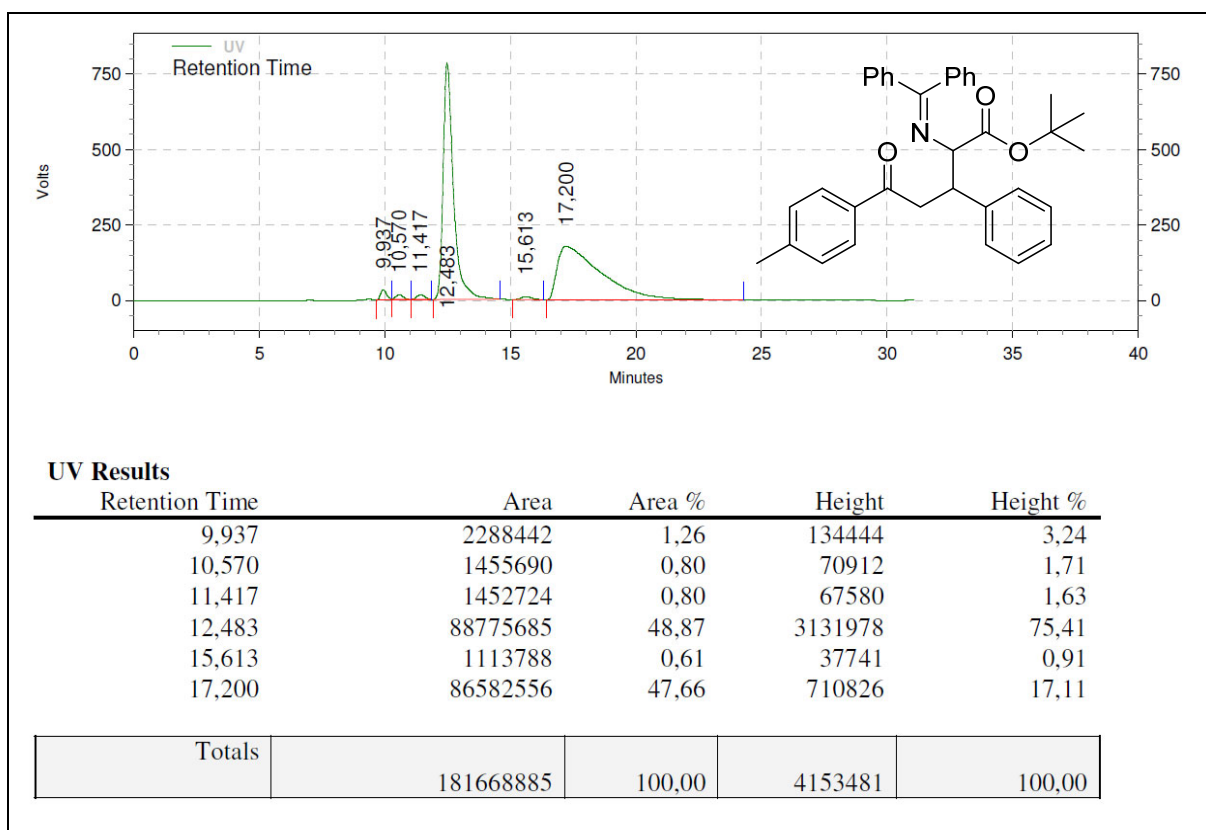
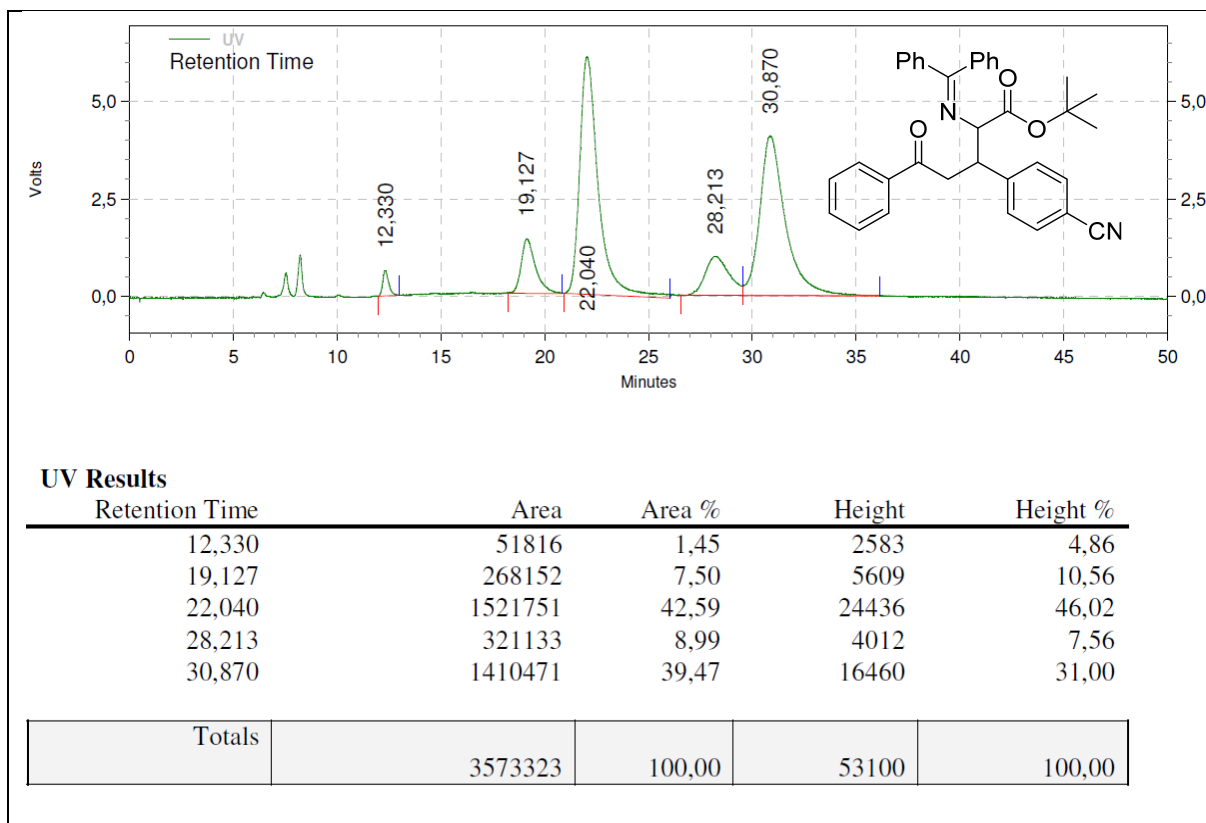
UV Results

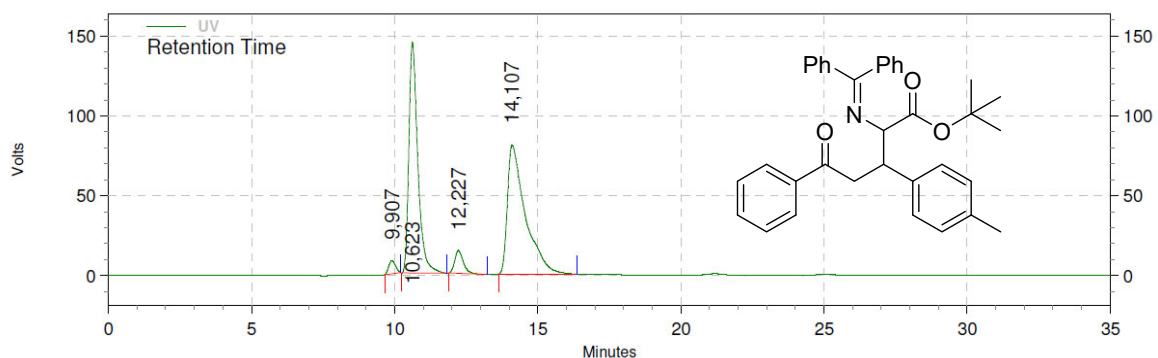
Retention Time	Area	Area %	Height	Height %
45,810	54188	0,07	488	0,10
52,667	145028	0,18	1065	0,21
70,820	40481504	49,66	315219	62,47
89,597	40829810	50,09	187816	37,22

Totals	81510530	100,00	504588	100,00
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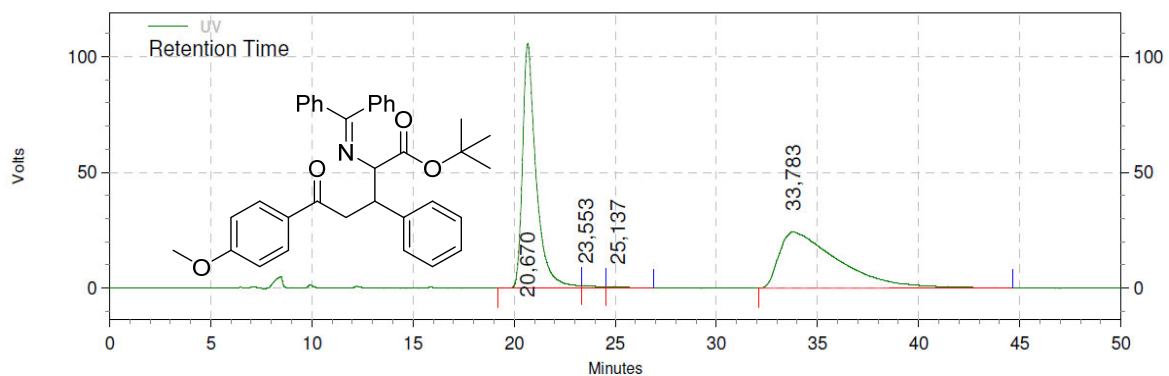






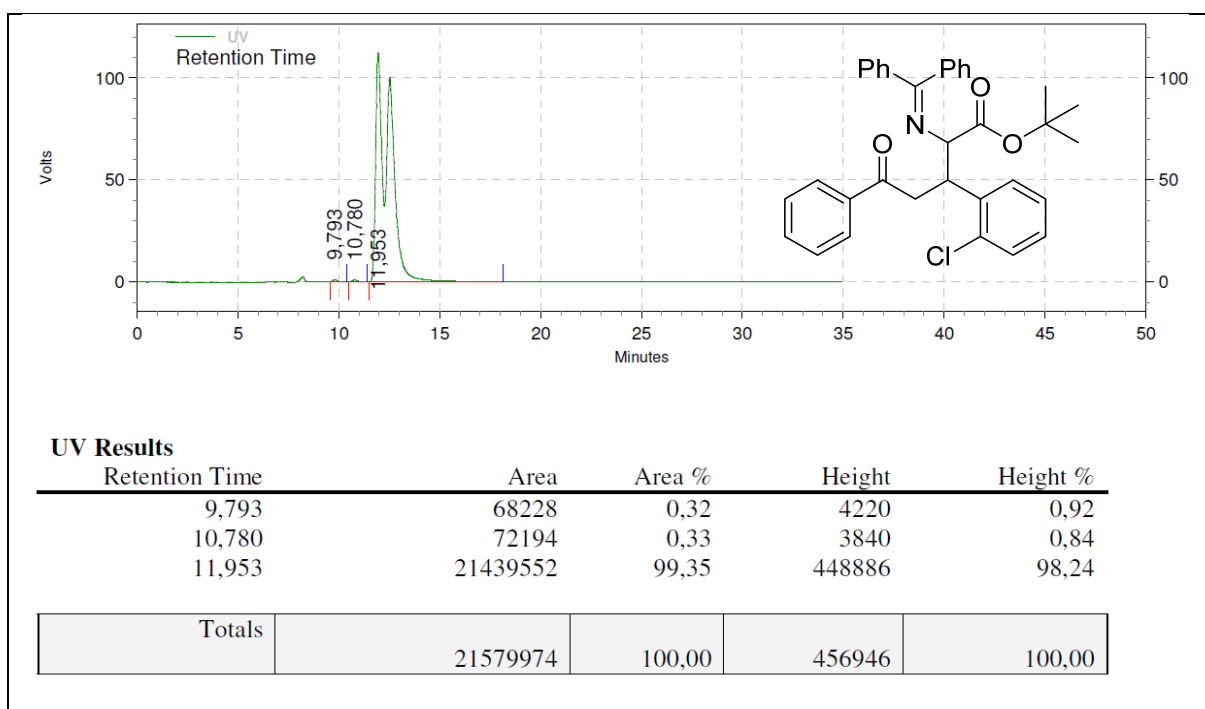
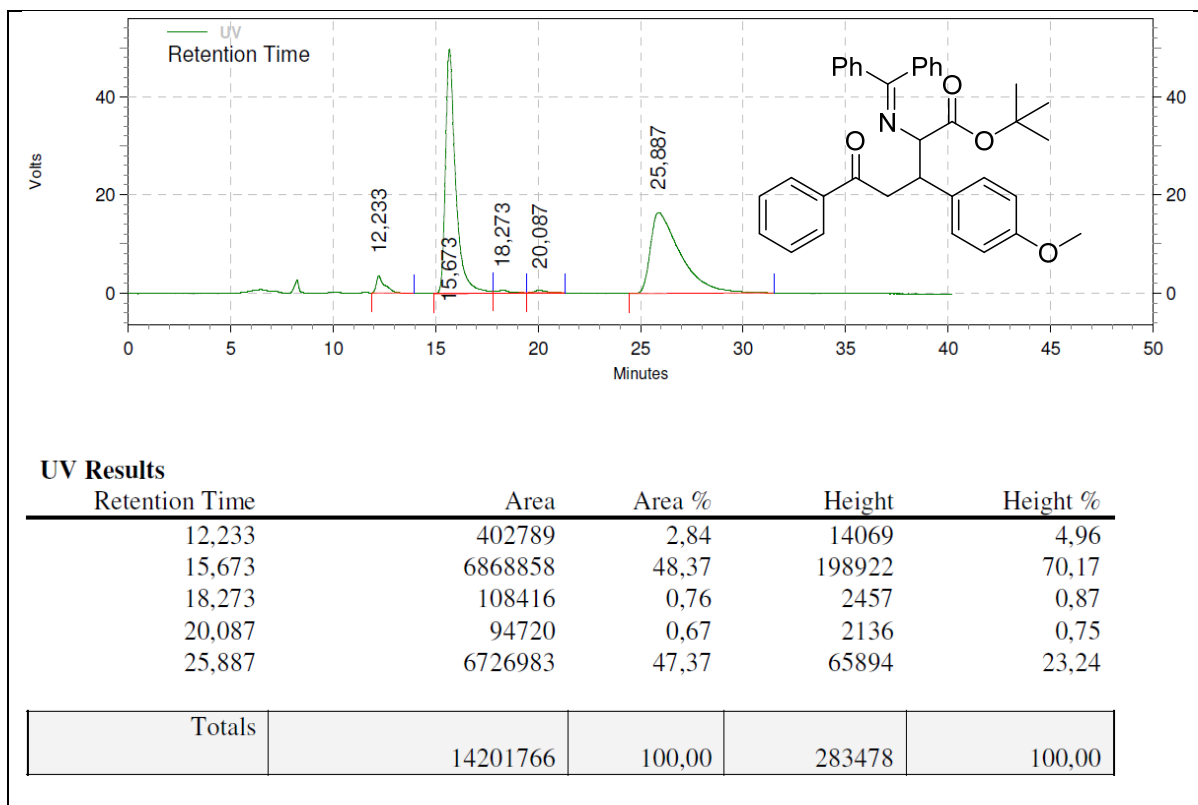
UV Results

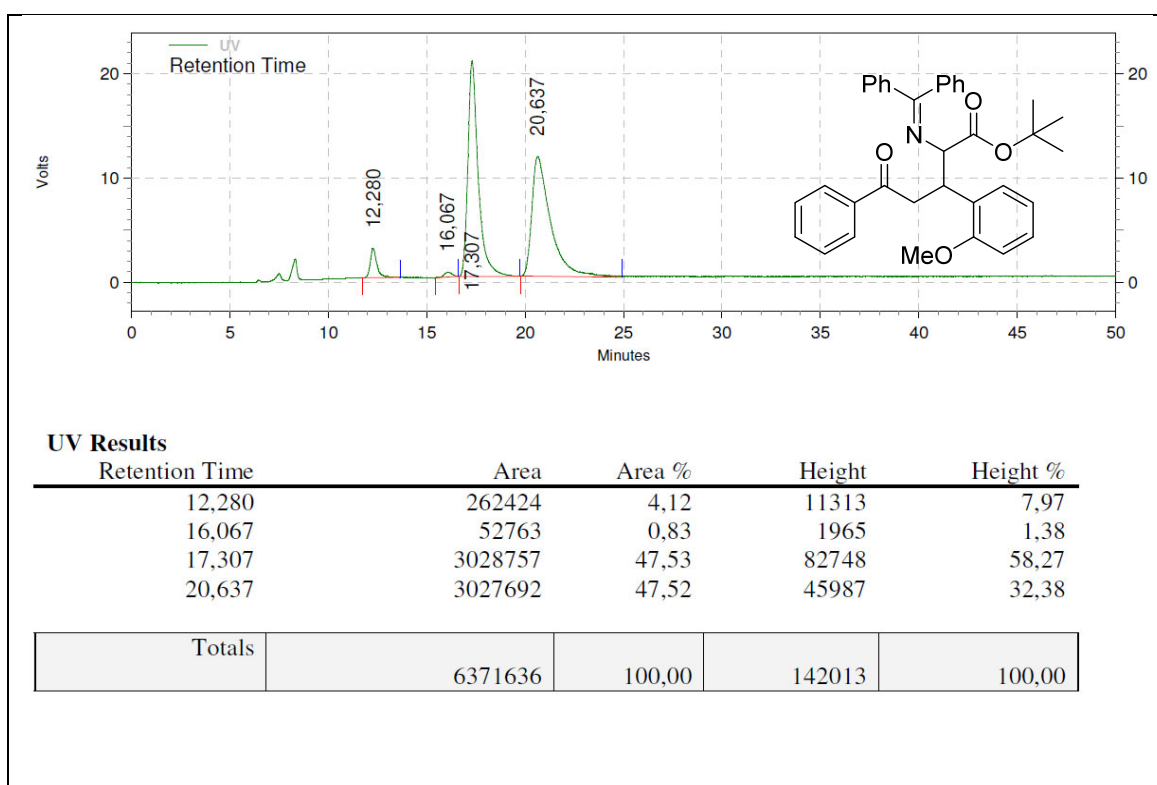
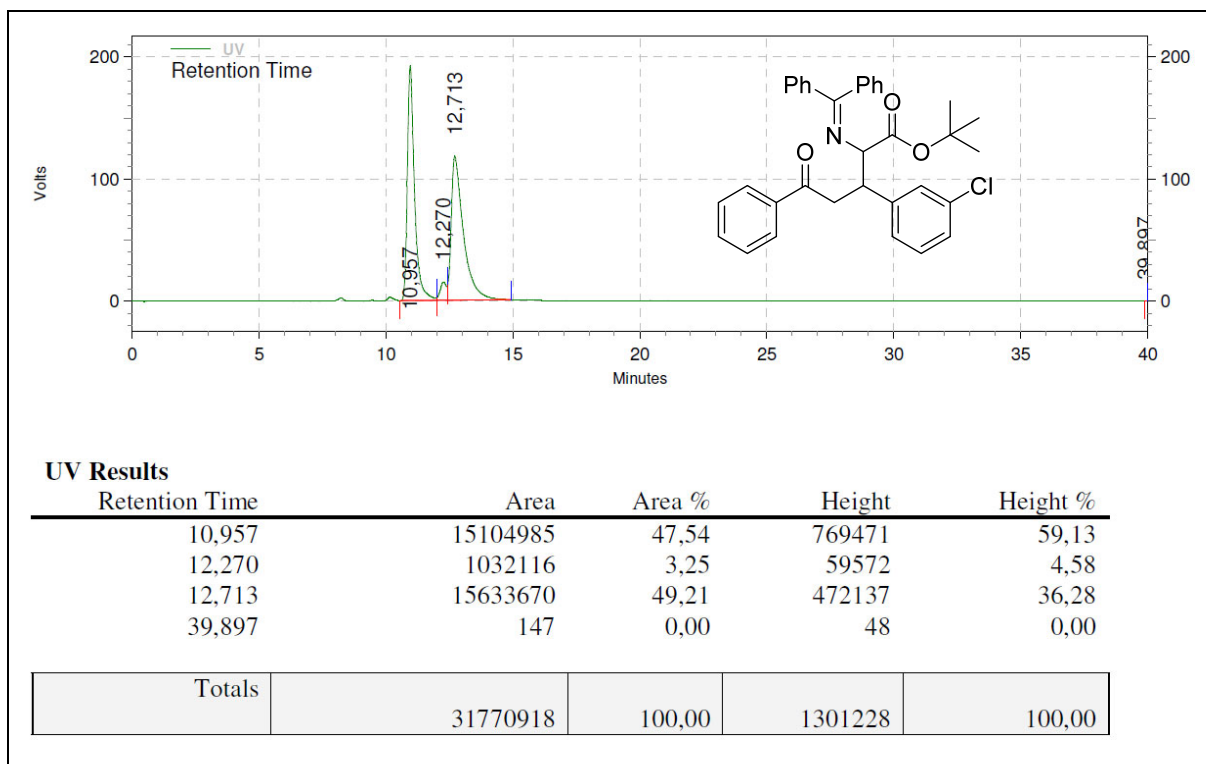
Retention Time	Area	Area %	Height	Height %
9,907	526123	1,86	32465	3,28
10,623	12537526	44,34	576840	58,27
12,227	1177590	4,16	57200	5,78
14,107	14032246	49,63	323482	32,68
Totals	28273485	100,00	989987	100,00

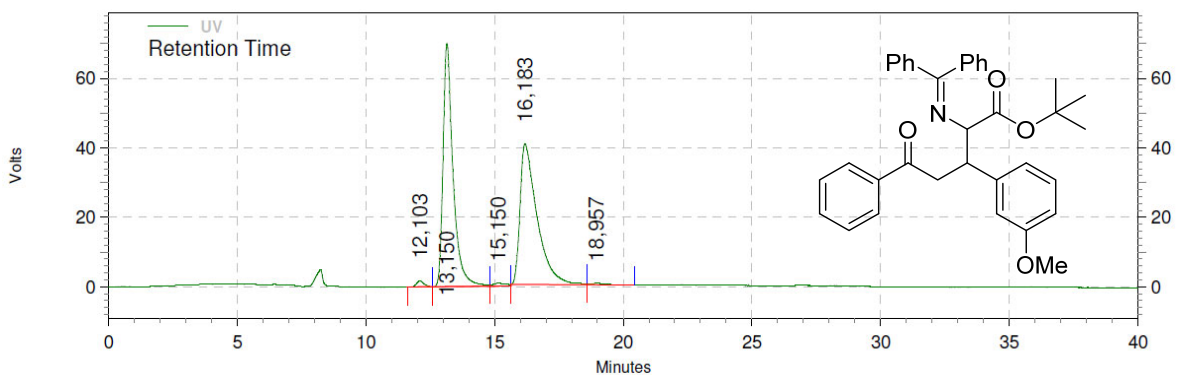


UV Results

Retention Time	Area	Area %	Height	Height %
20,670	19715310	49,60	422294	80,42
23,553	197694	0,50	3739	0,71
25,137	125601	0,32	2023	0,39
33,783	19706039	49,58	97080	18,49
Totals	39744644	100,00	525136	100,00

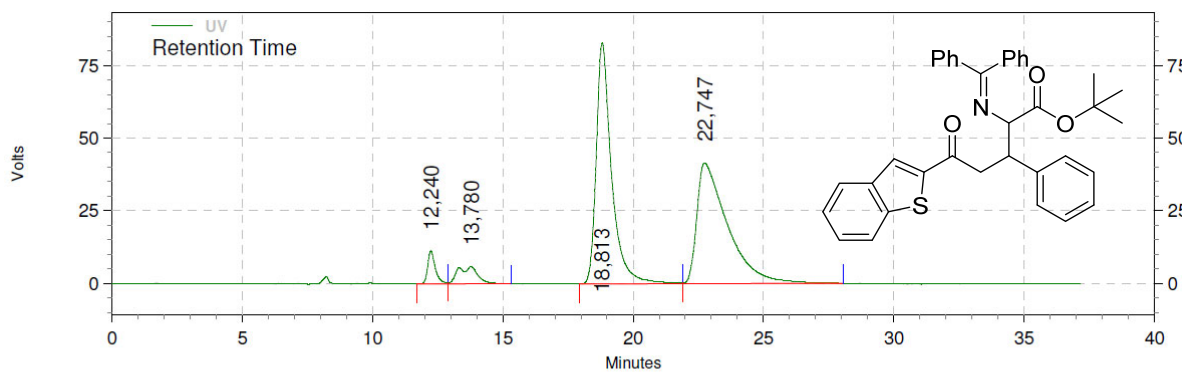






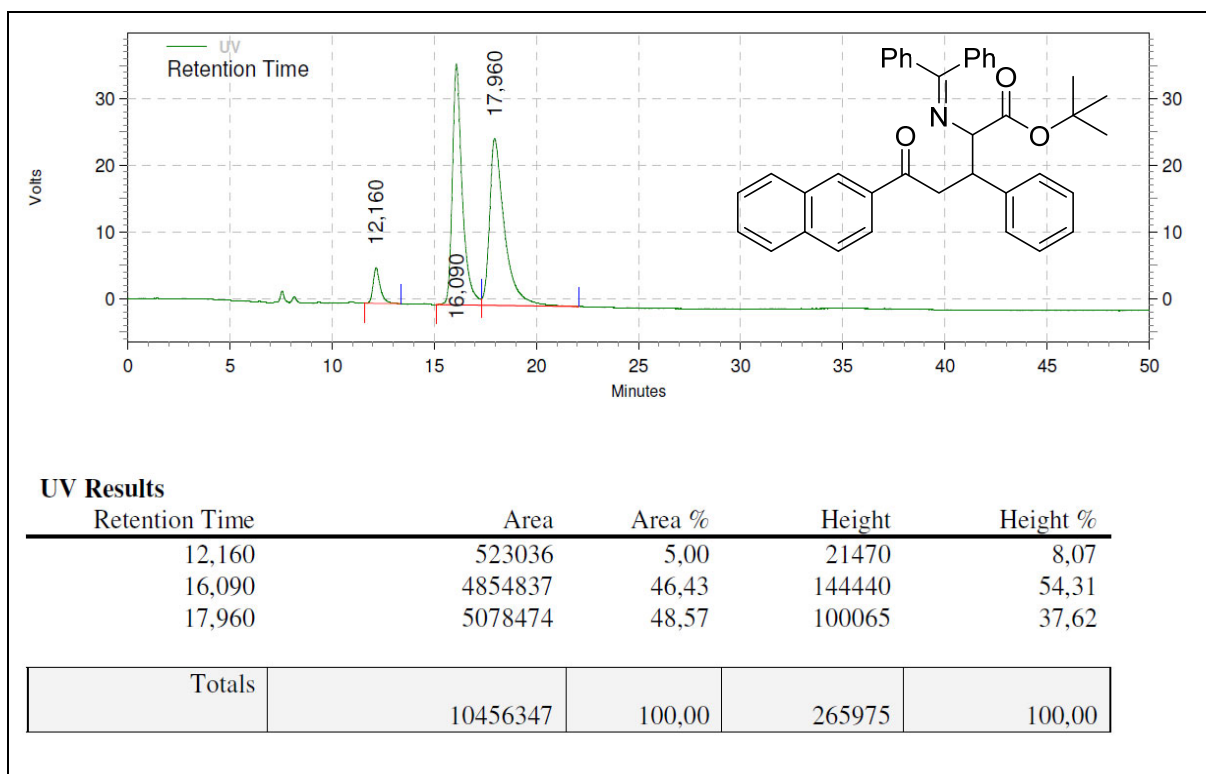
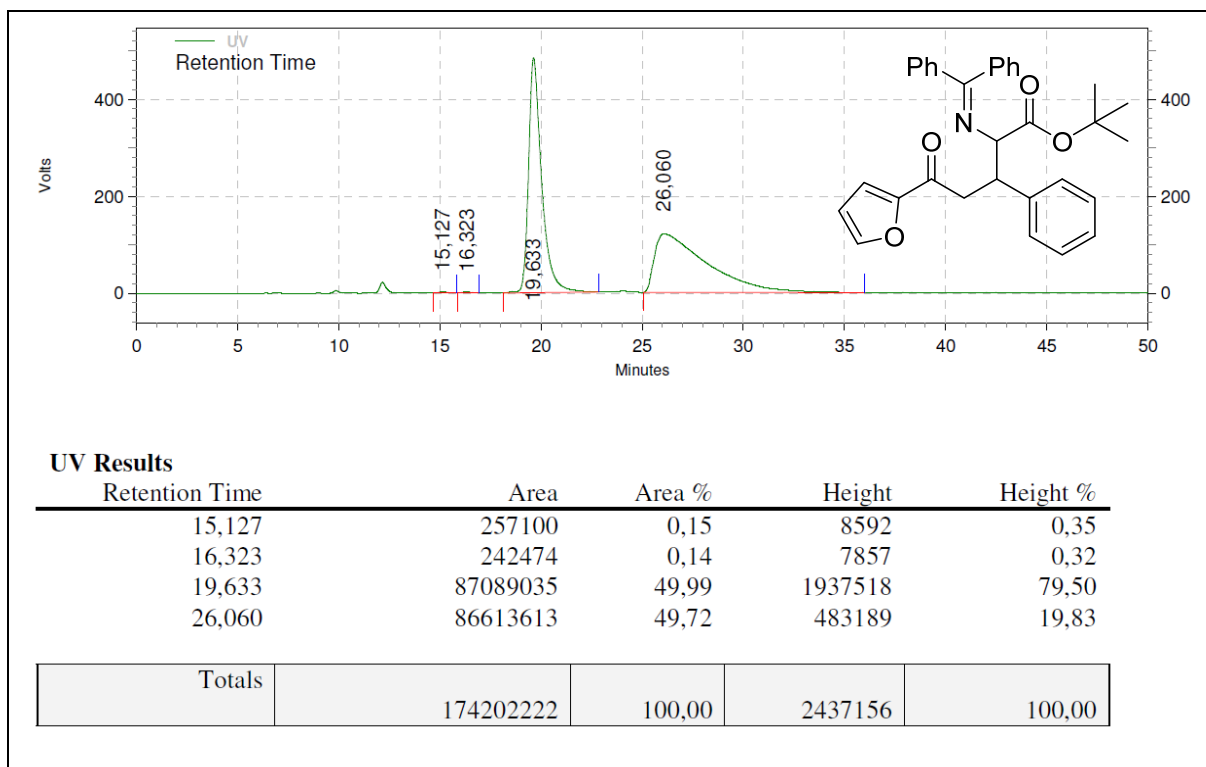
UV Results

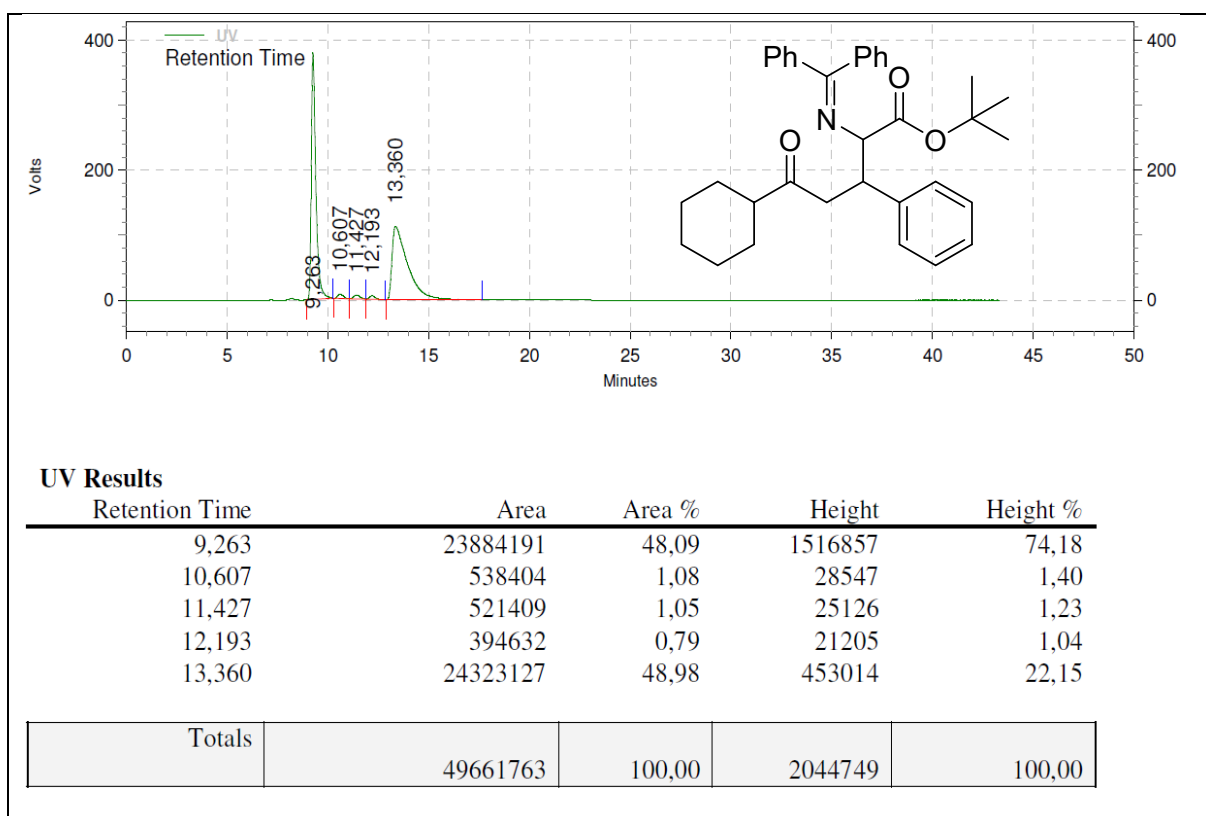
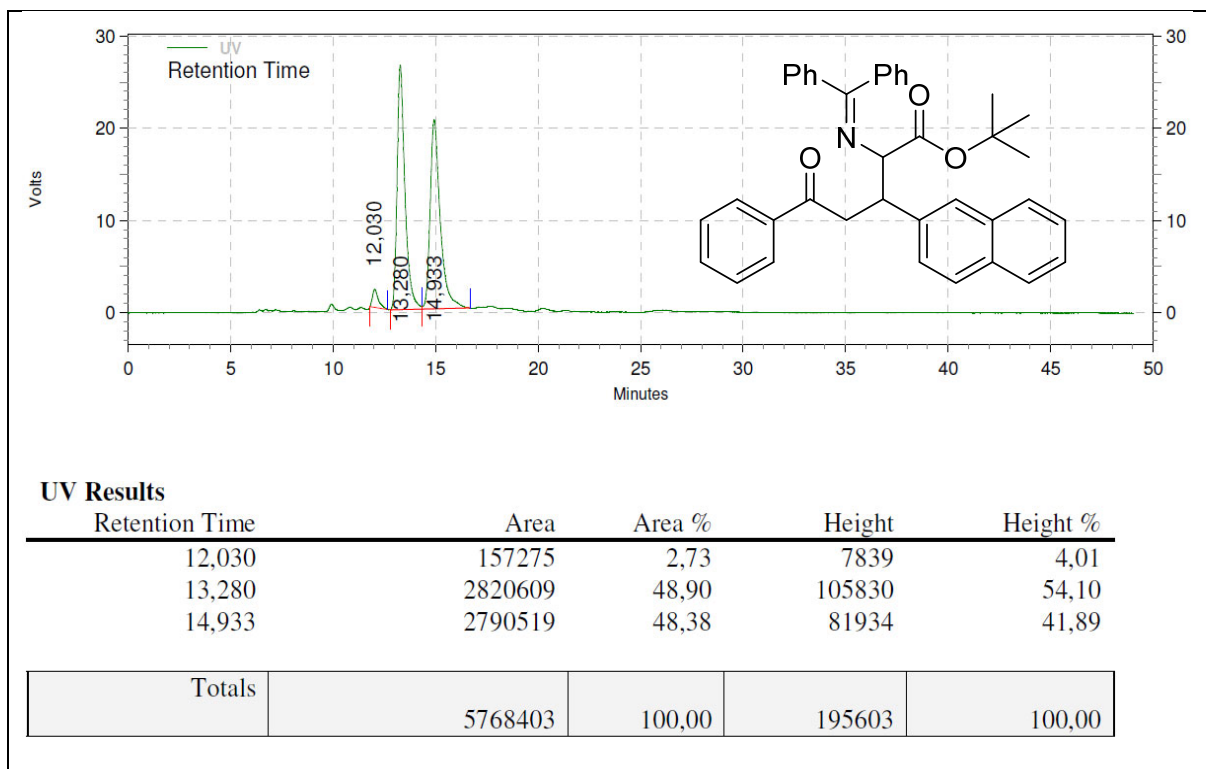
Retention Time	Area	Area %	Height	Height %
12,103	135840	0,88	6937	1,53
13,150	7562044	49,21	280088	61,69
15,150	119198	0,78	3266	0,72
16,183	7471148	48,61	162085	35,70
18,957	80072	0,52	1628	0,36
Totals	15368302	100,00	454004	100,00

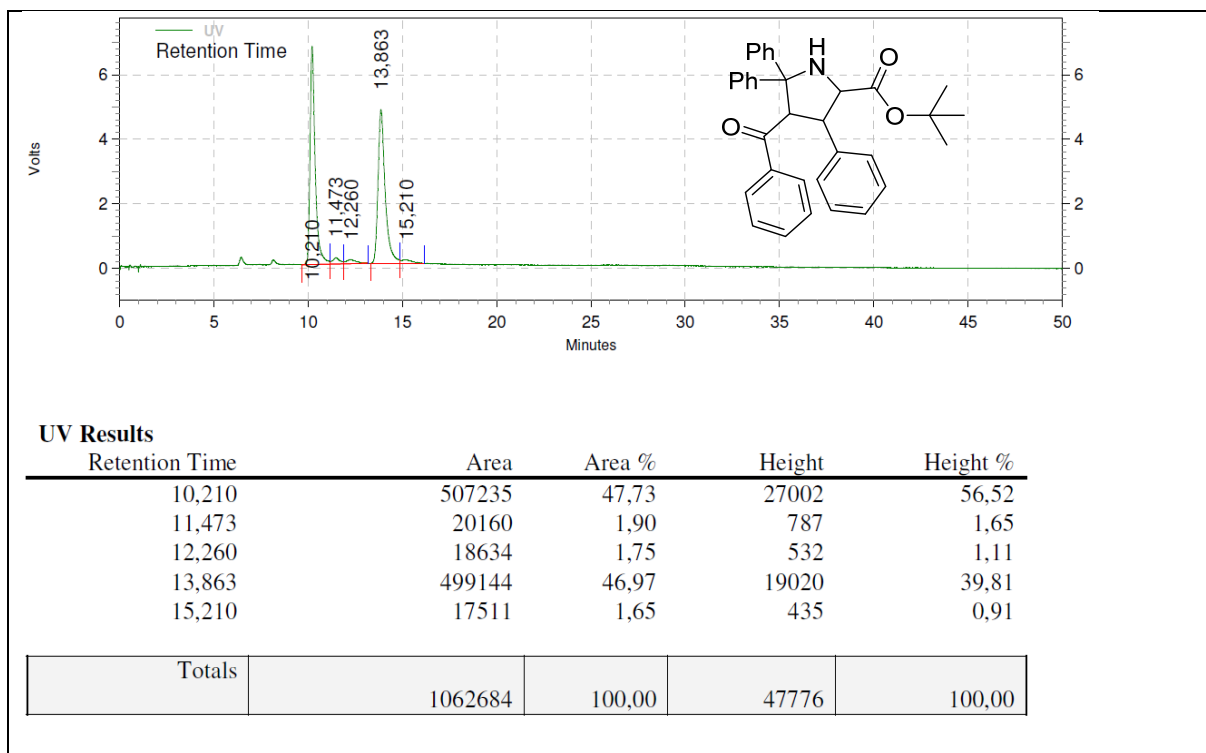
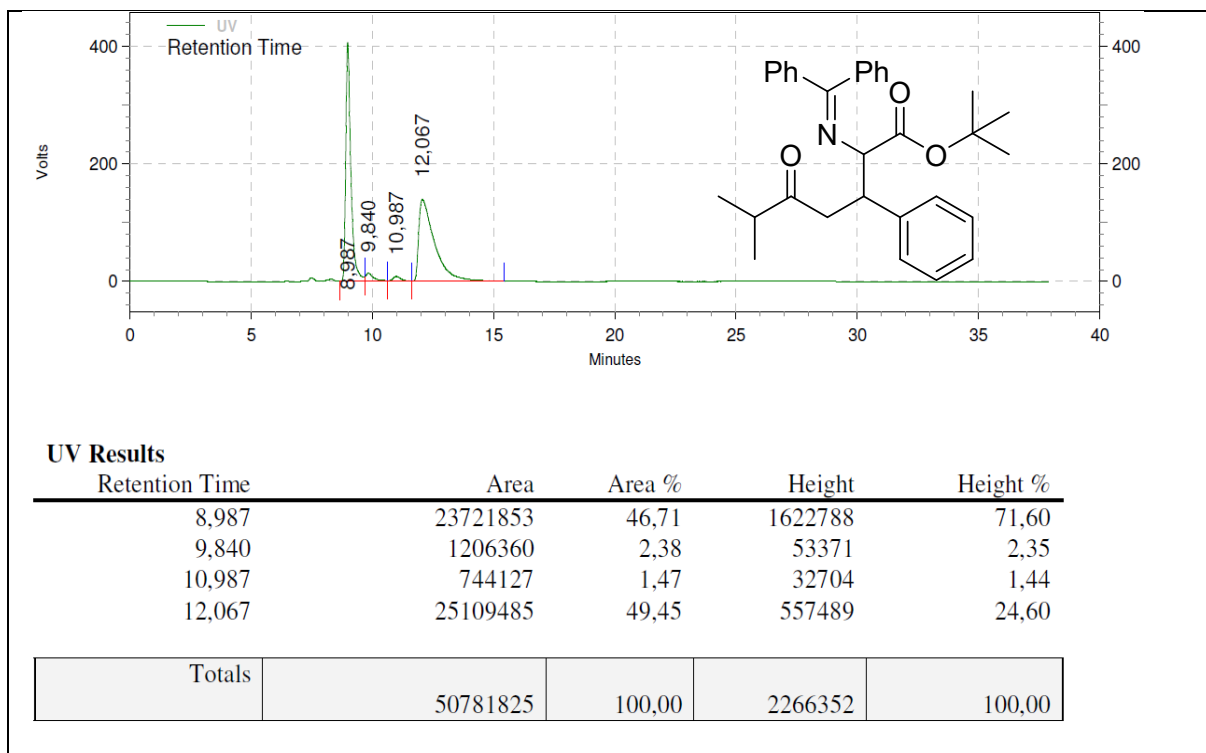


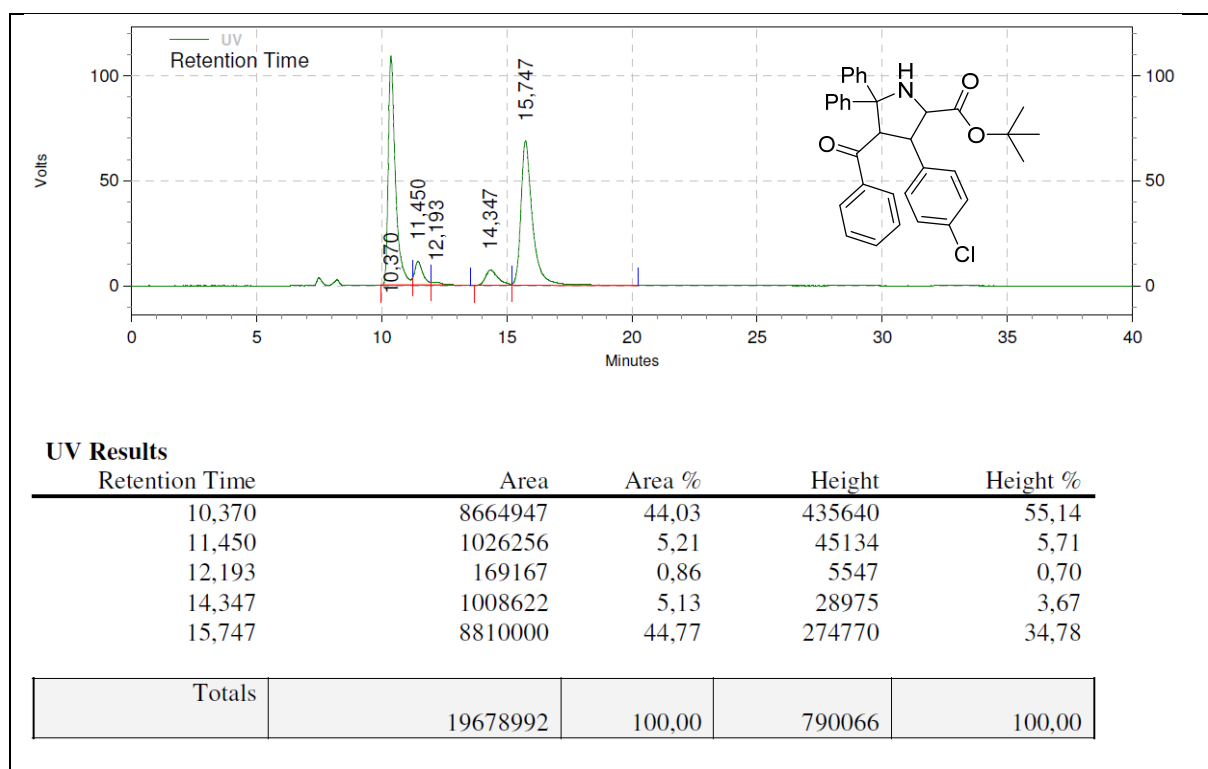
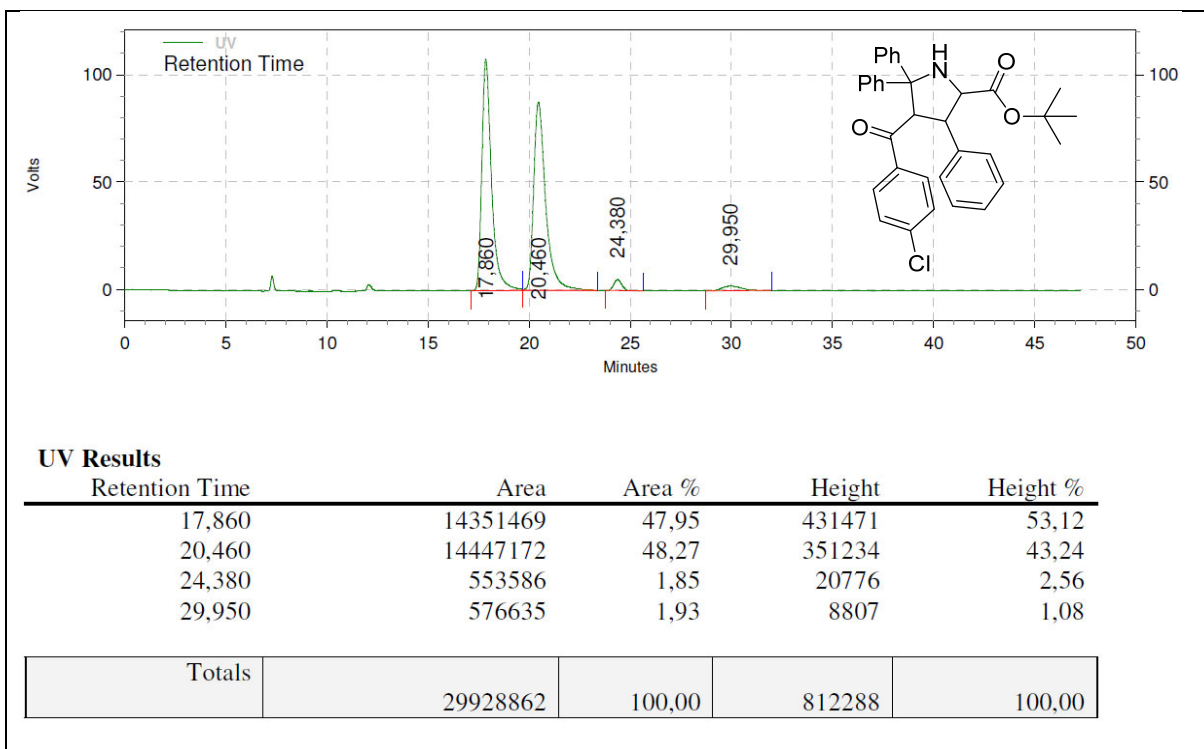
UV Results

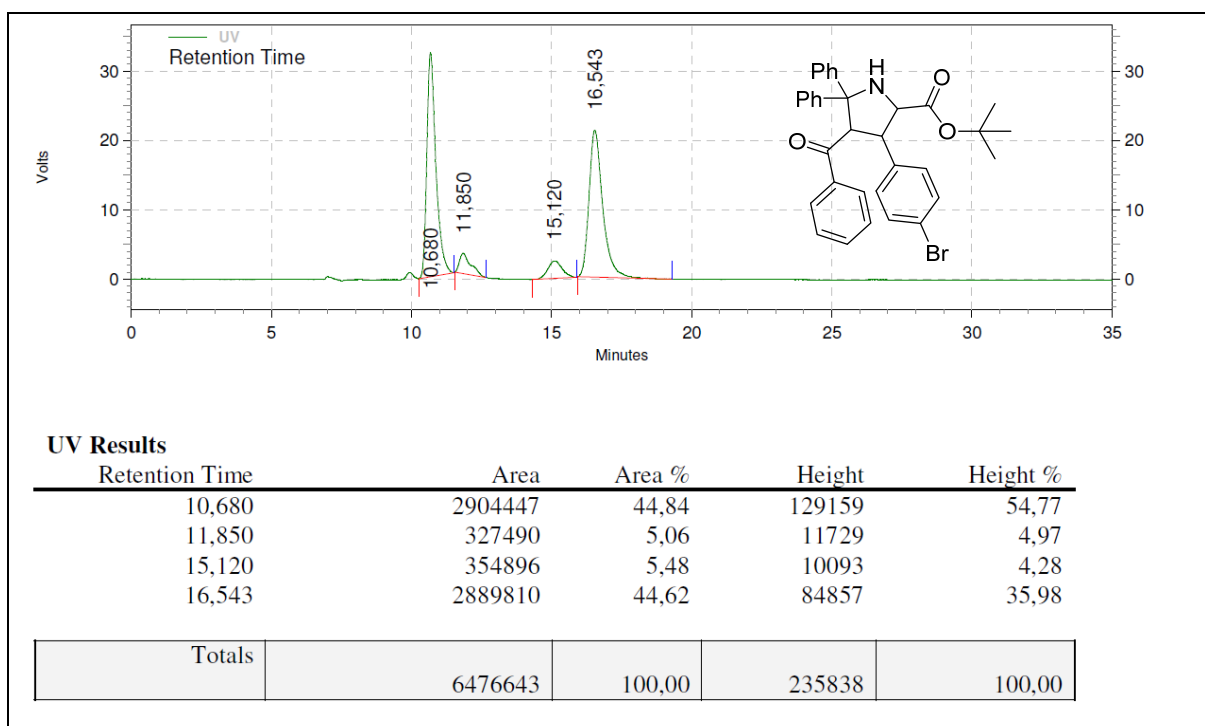
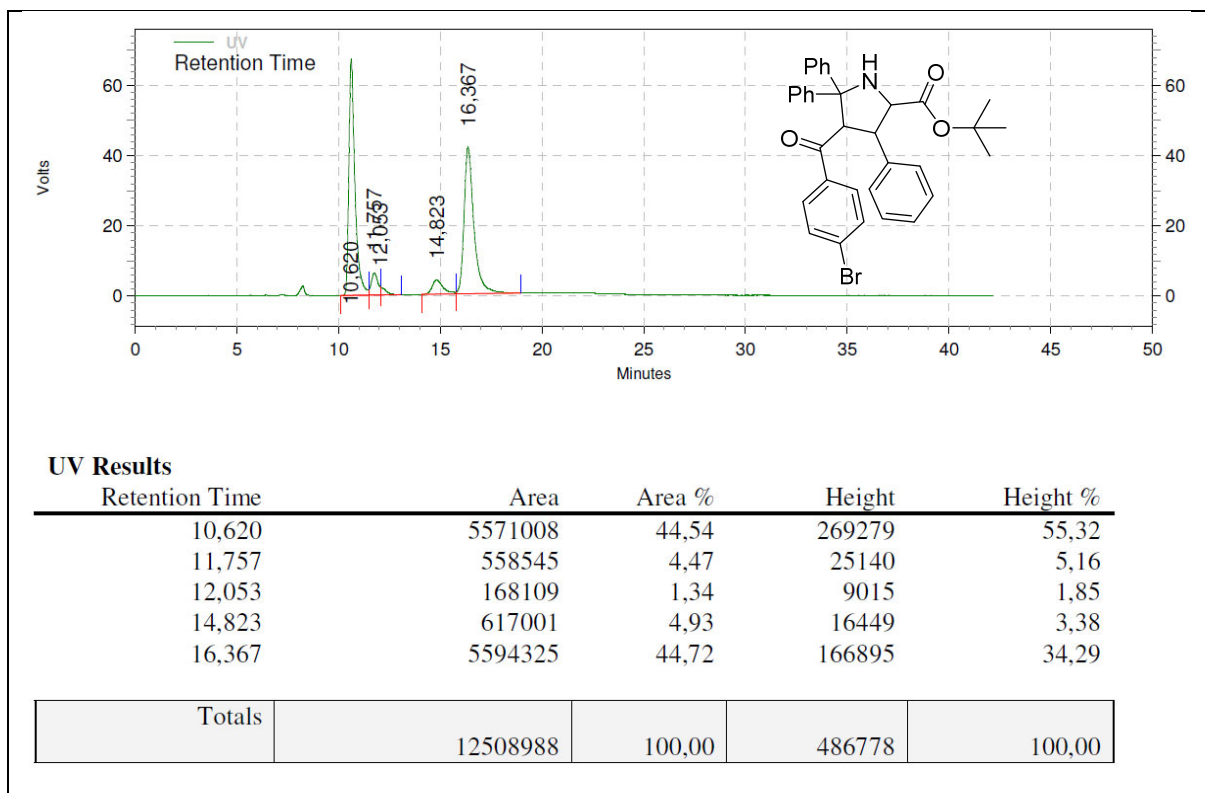
Retention Time	Area	Area %	Height	Height %
12,240	924325	3,14	45422	8,00
13,780	1251680	4,26	23690	4,17
18,813	13624925	46,34	332148	58,52
22,747	13604194	46,26	166343	29,31
Totals	29405124	100,00	567603	100,00

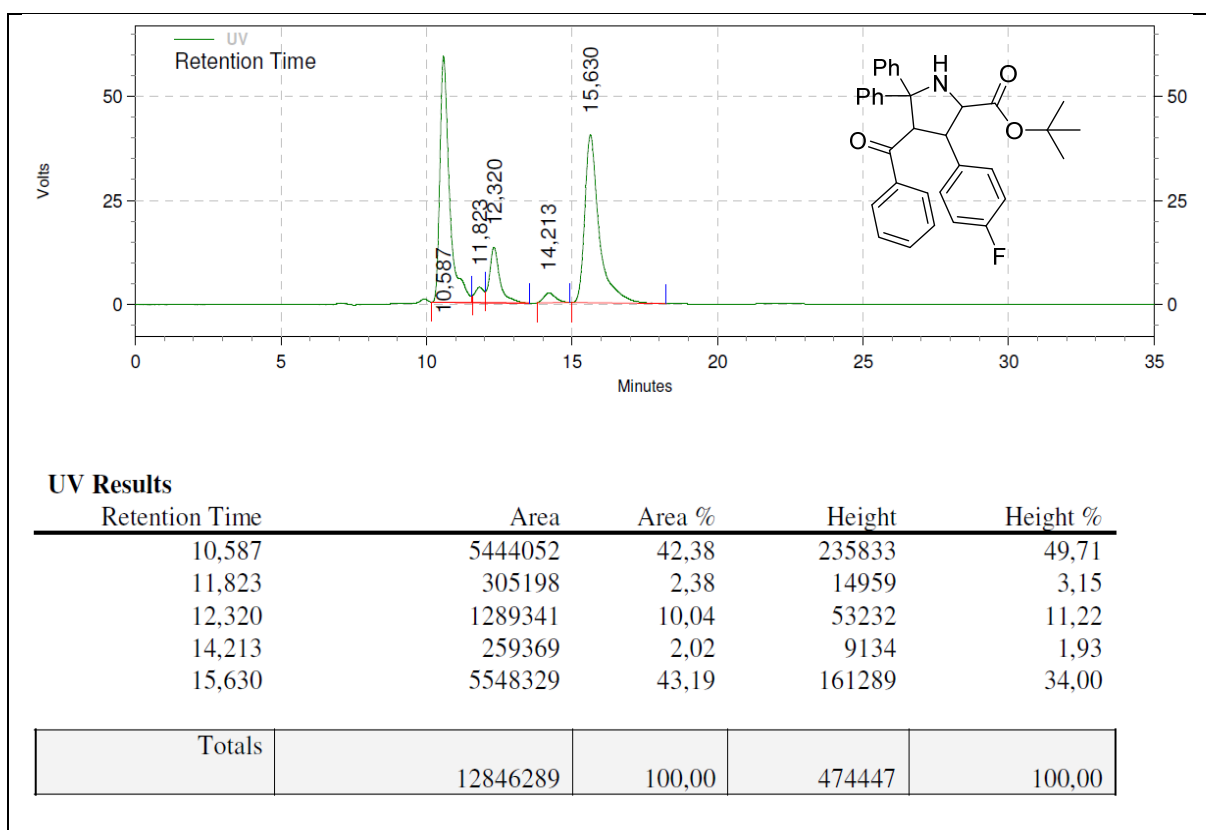
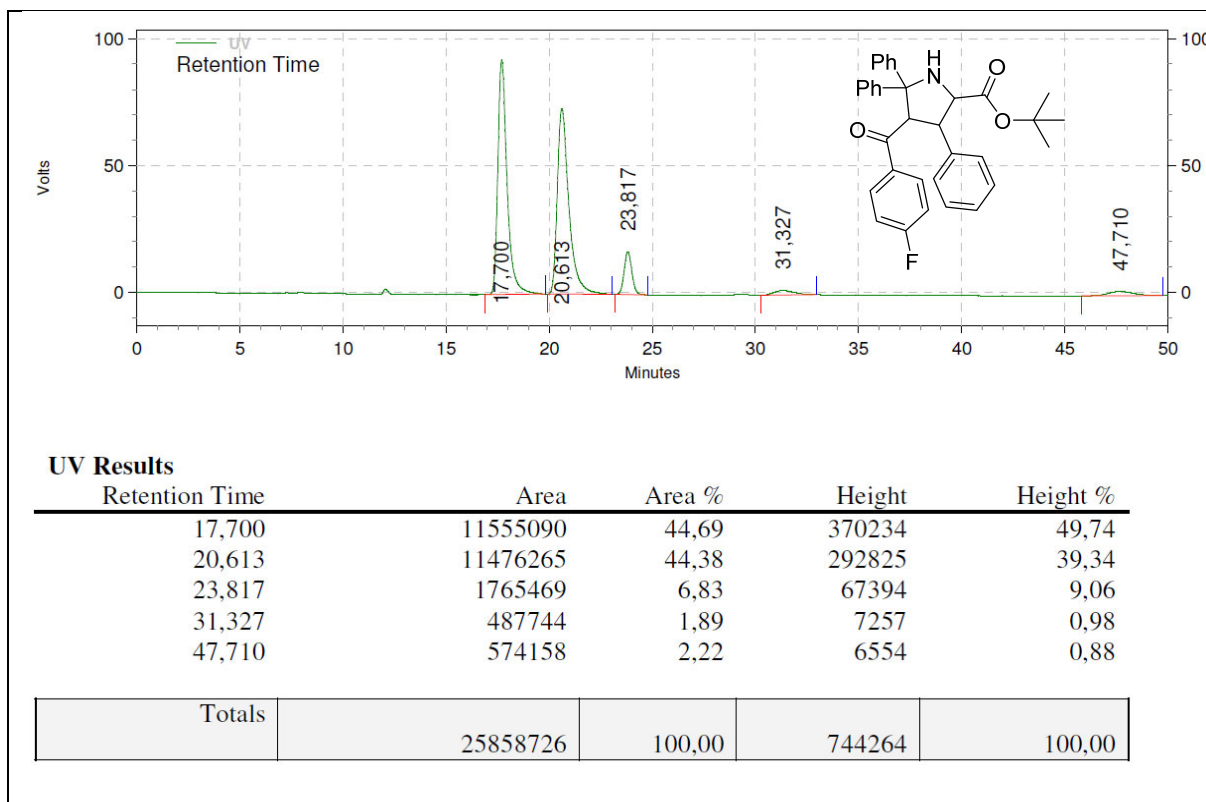


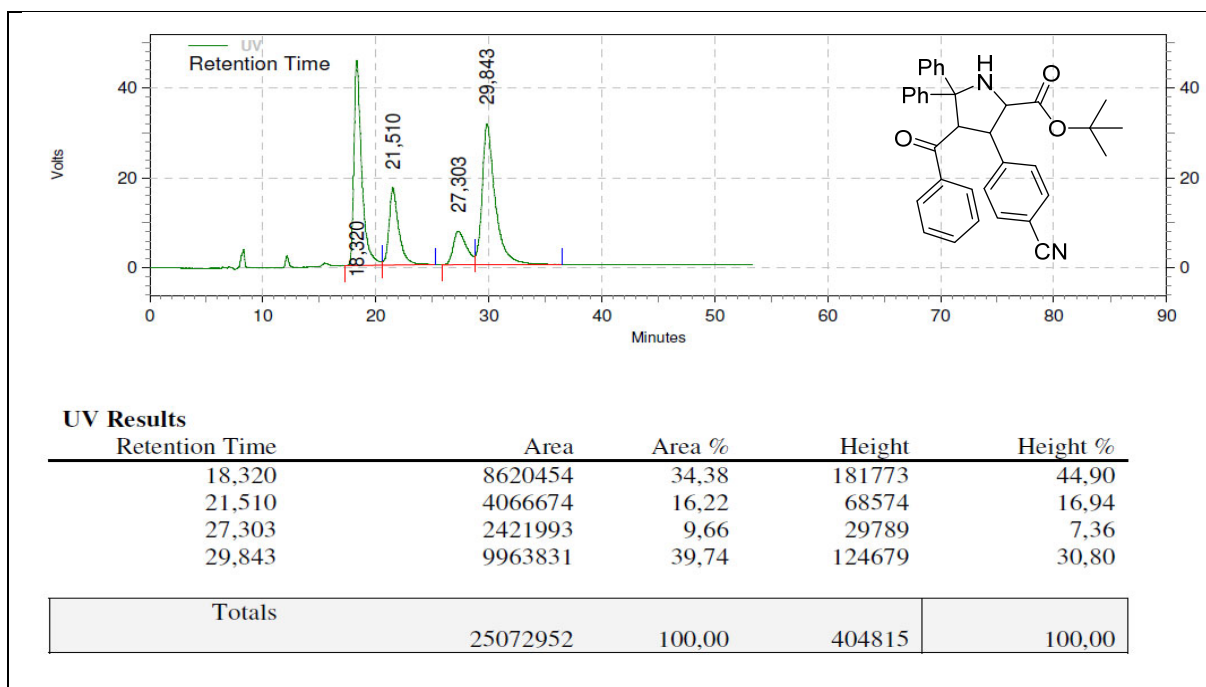
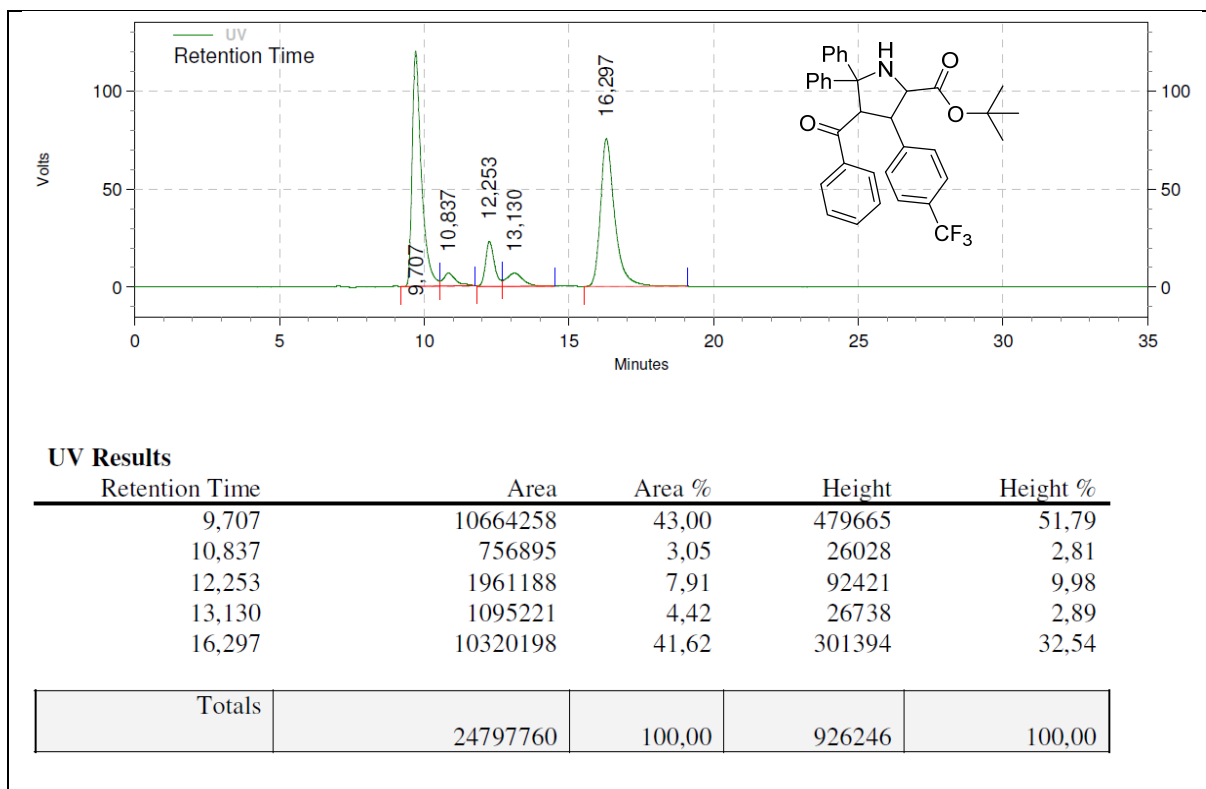


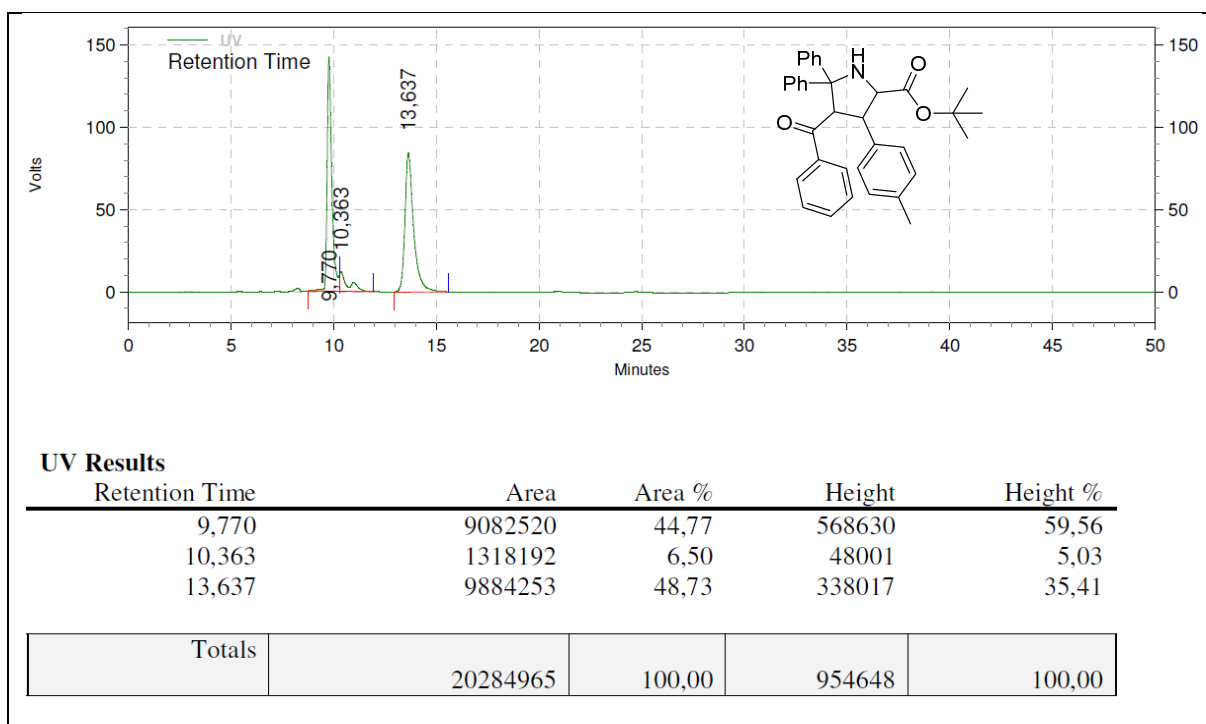
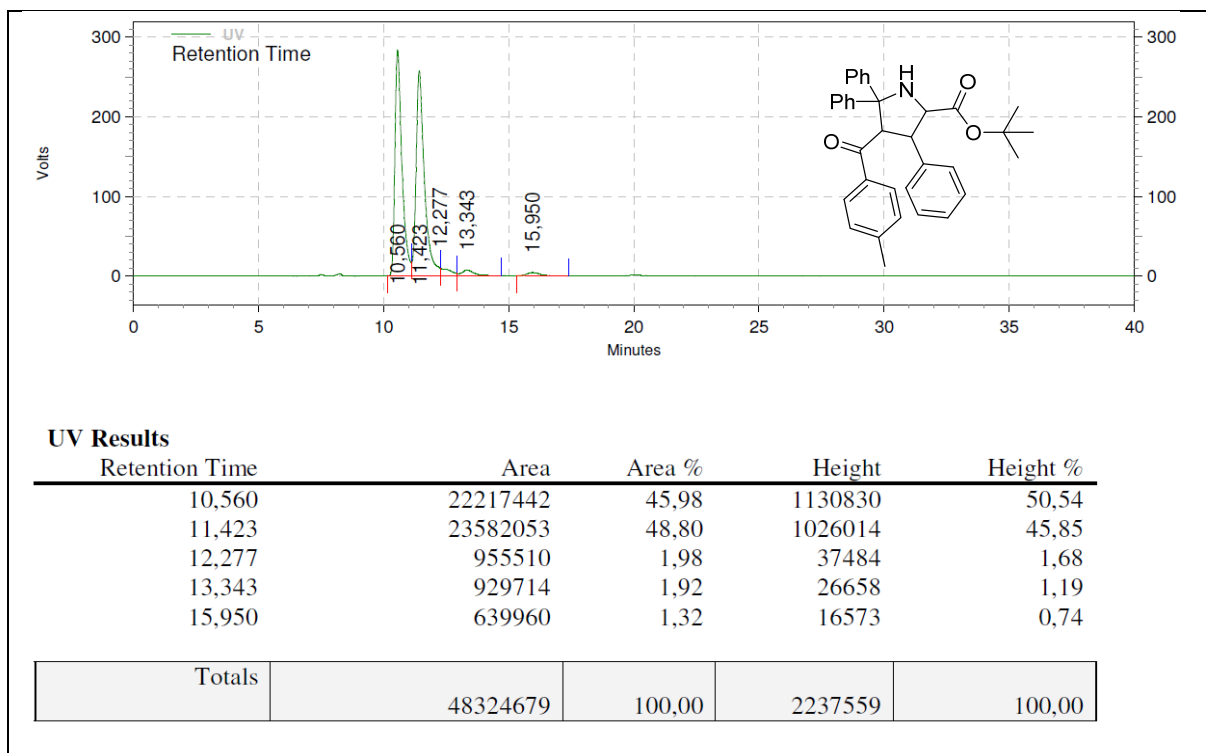


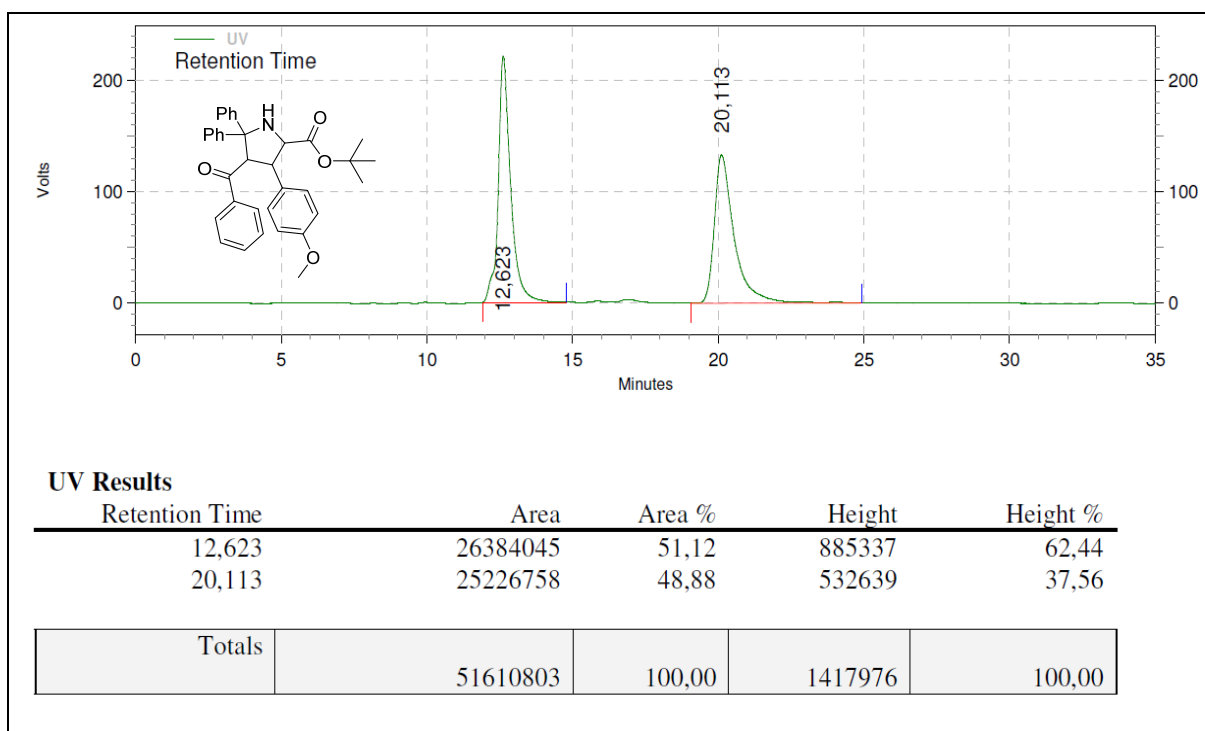
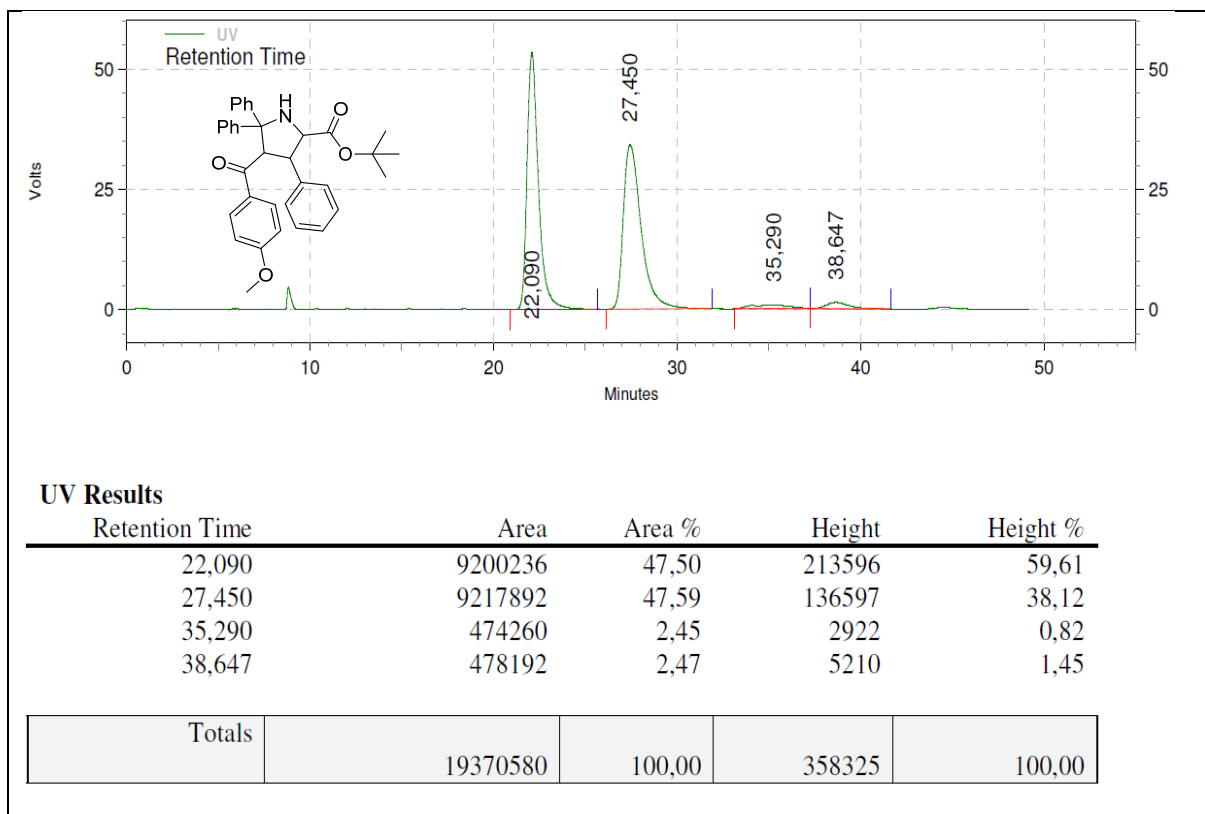


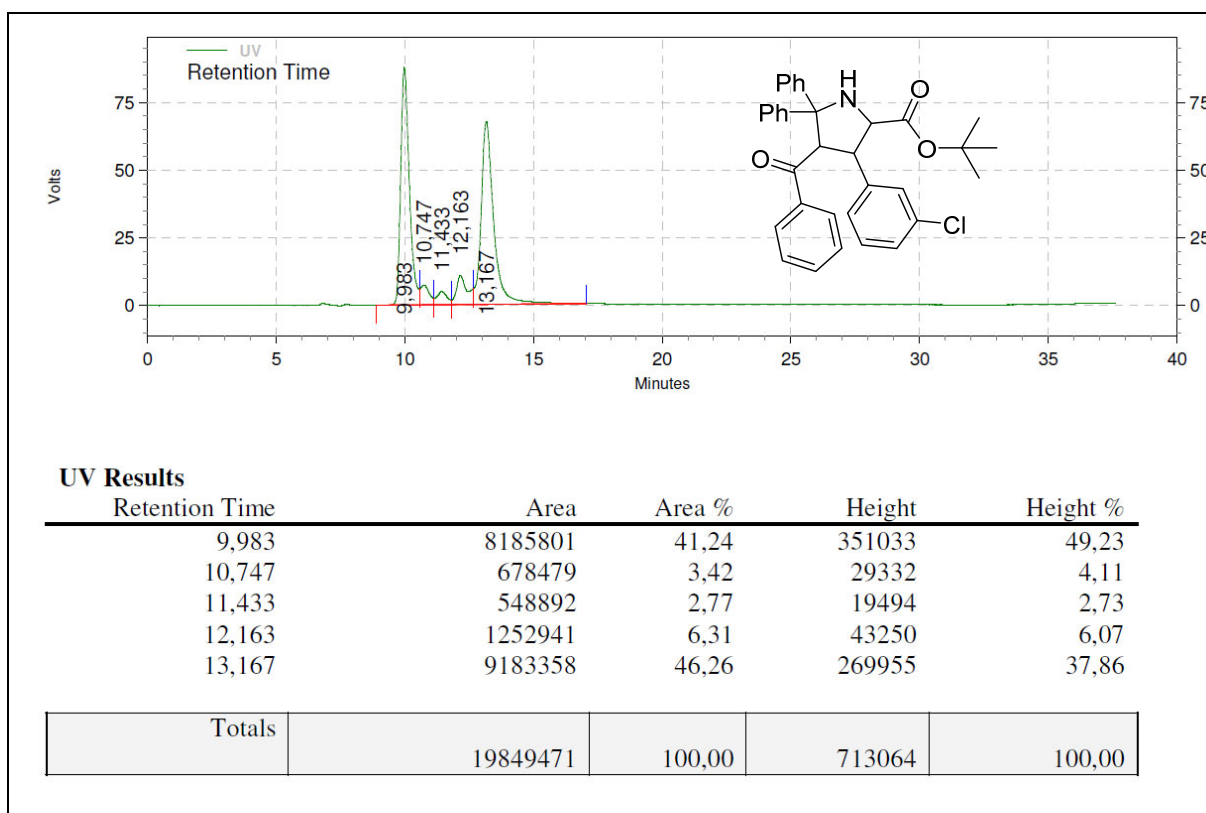
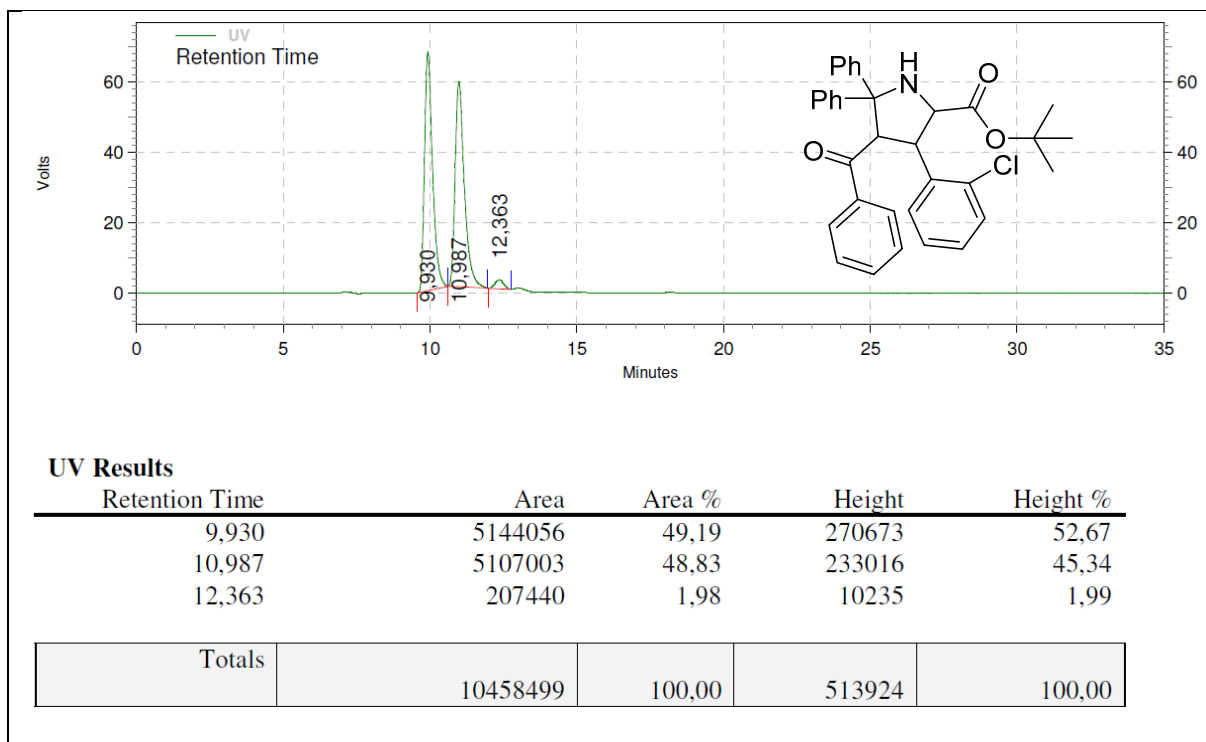


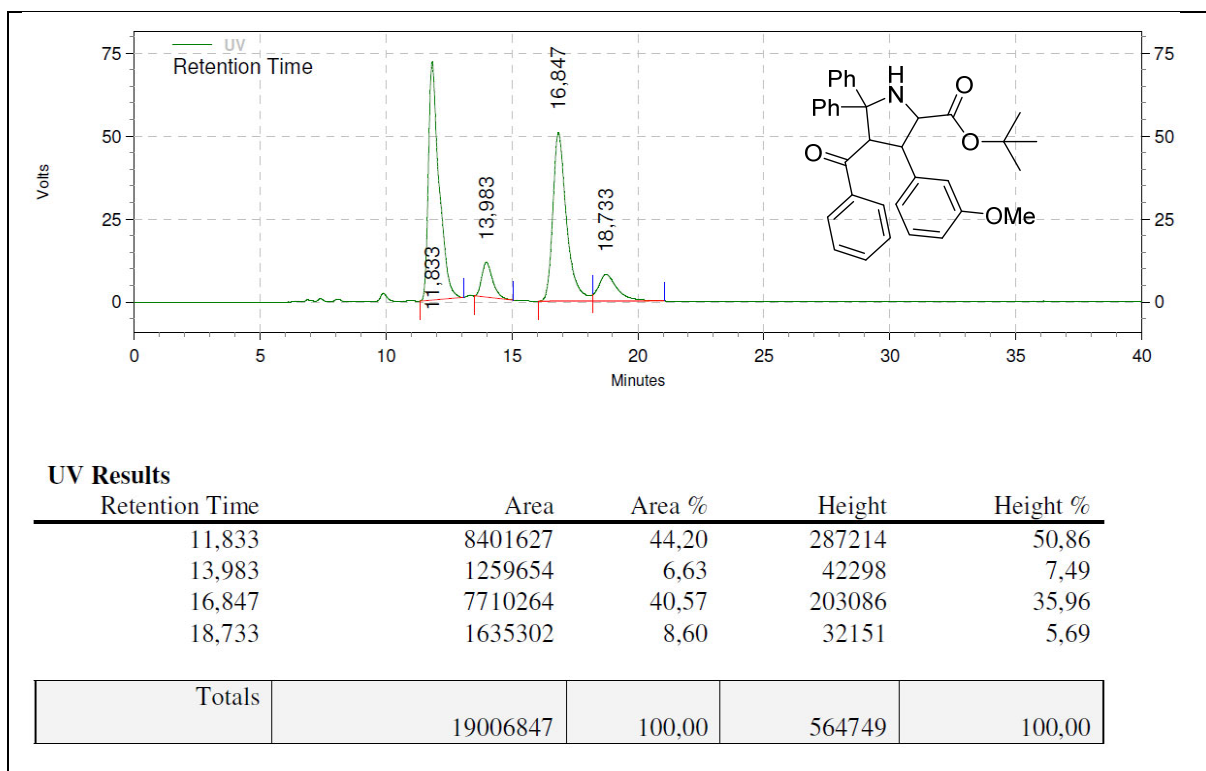
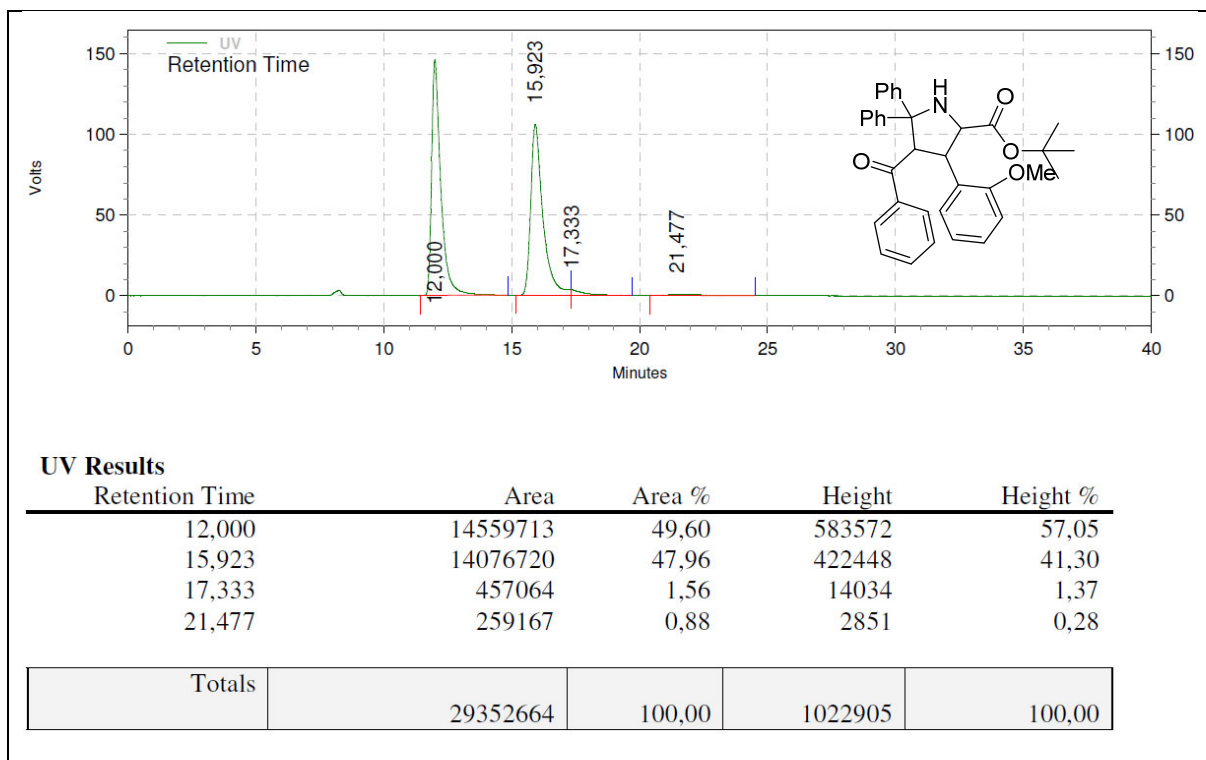


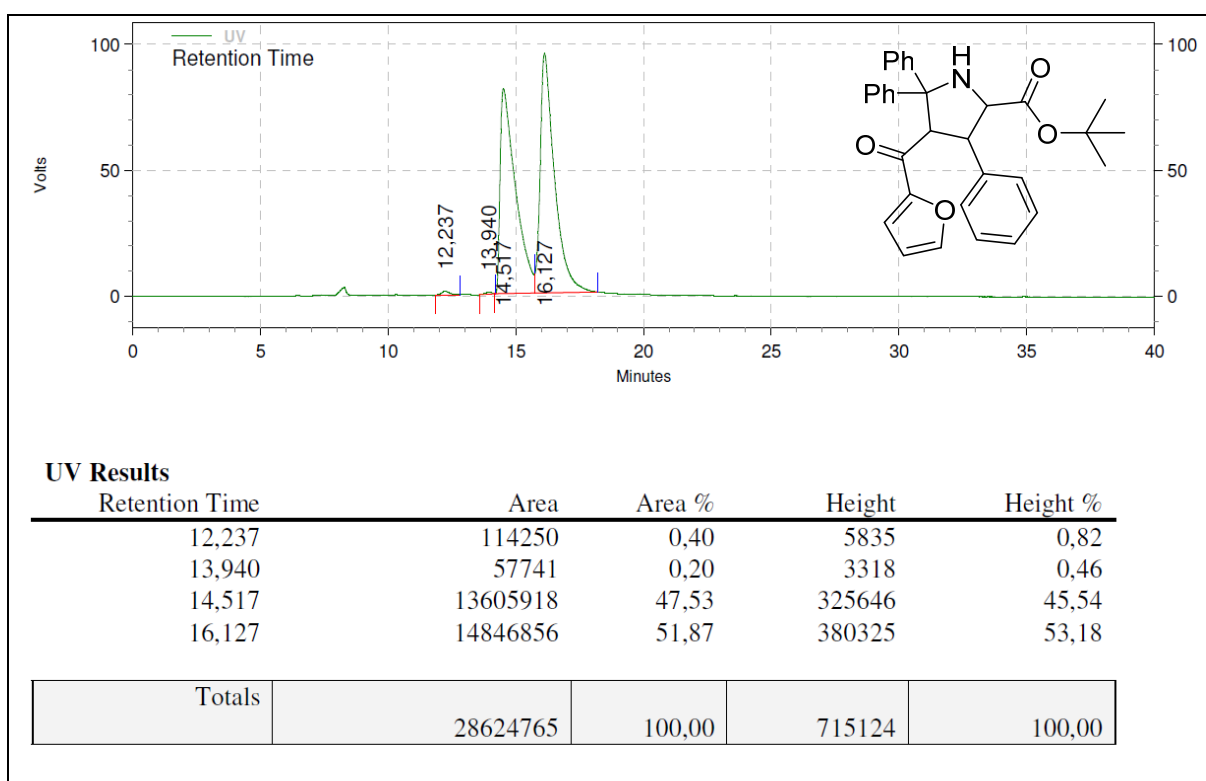
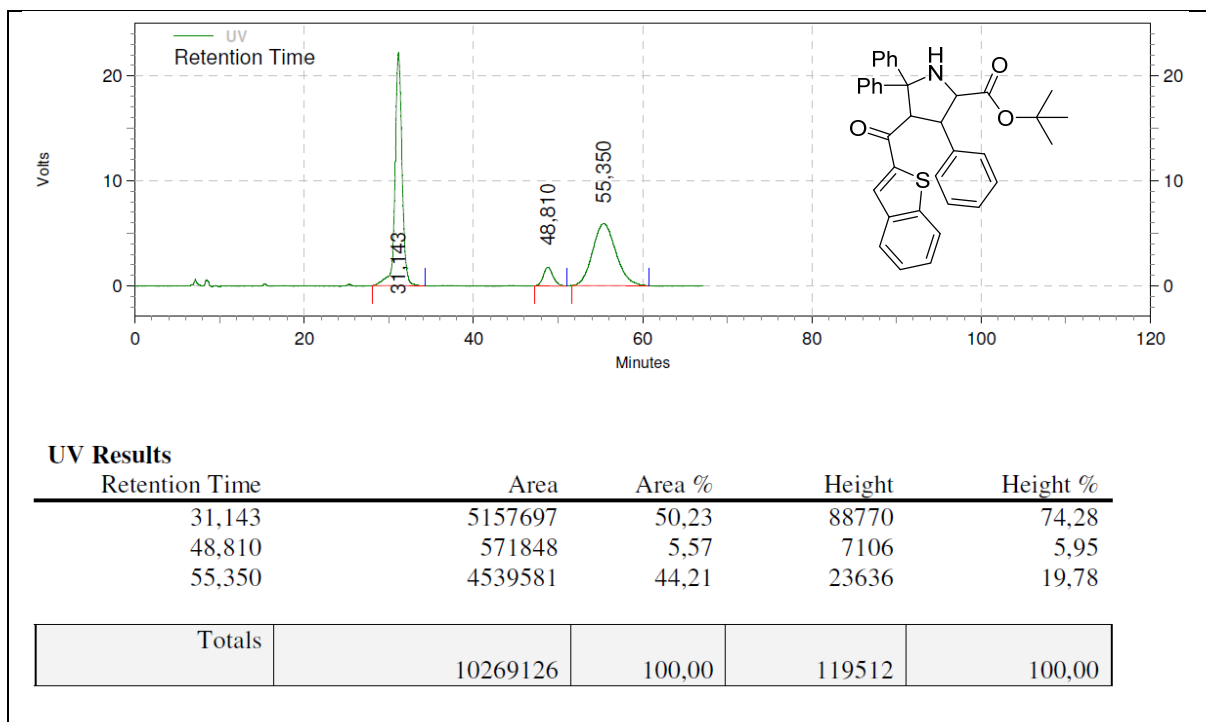


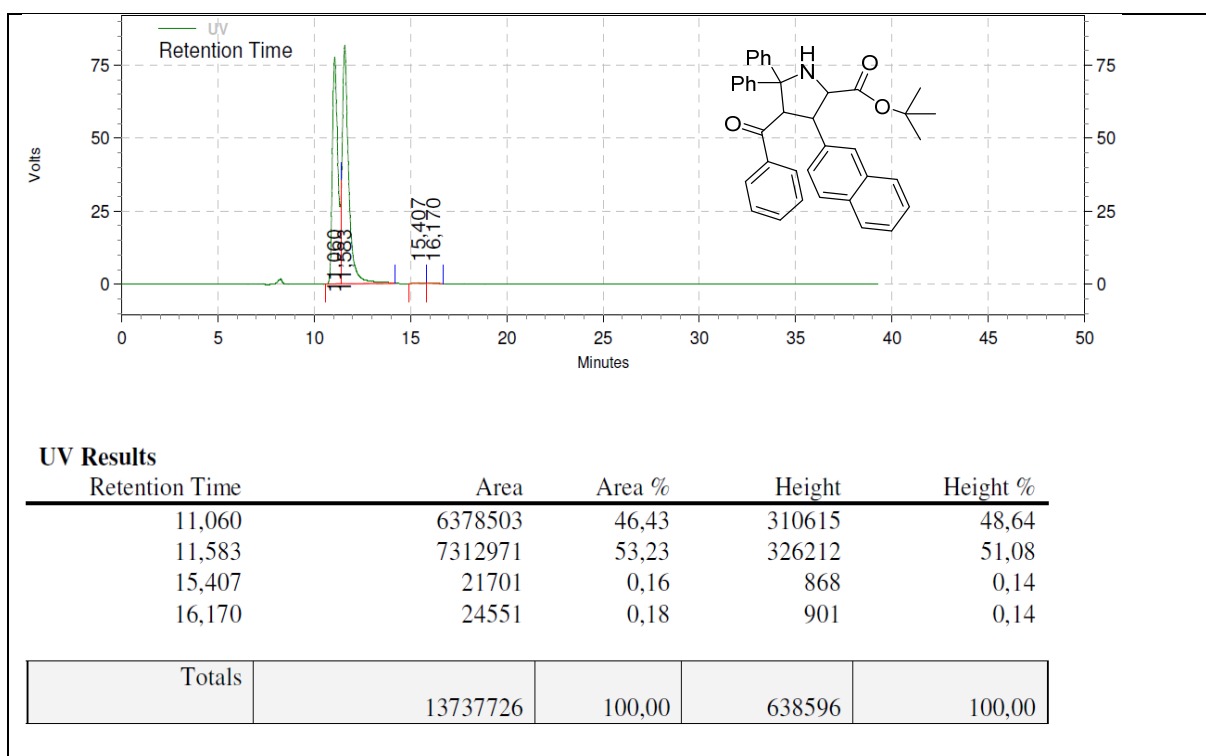
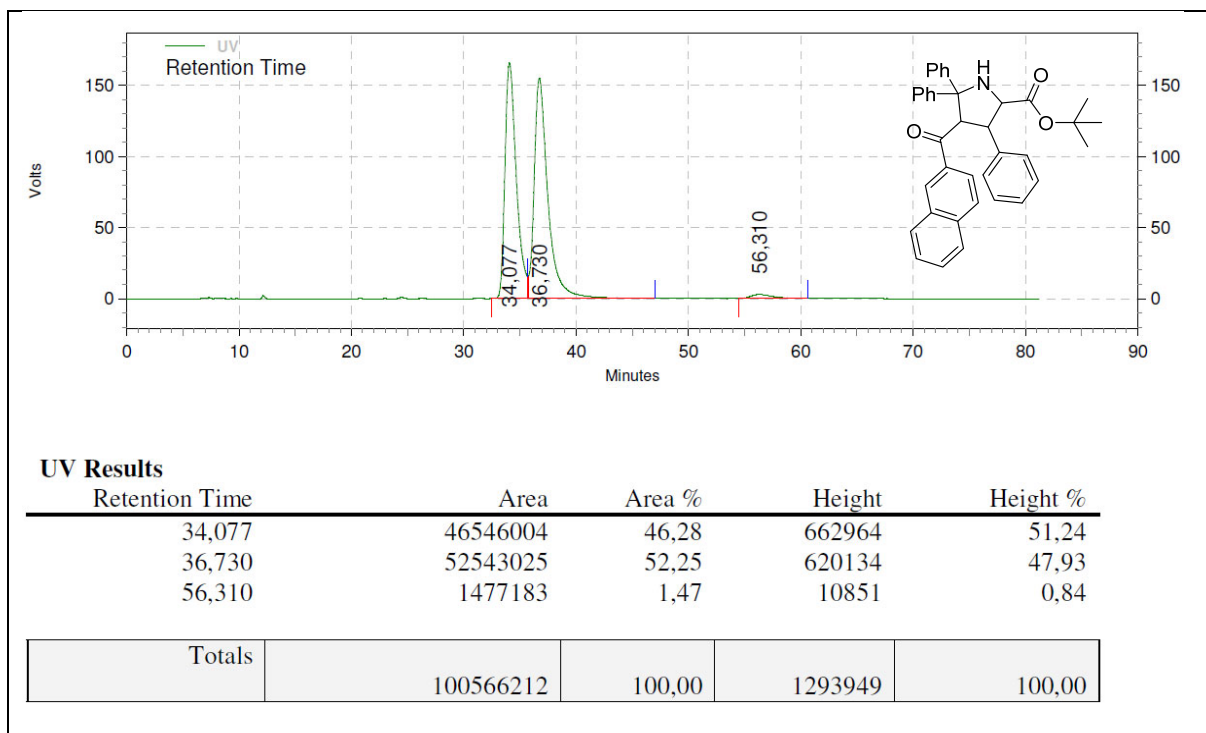












5. References

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