

Accessing Alternative Reaction Pathways of the Intermolecular Condensation between Homo-Propargyl Alcohols and Terminal Alkynes through Divergent Gold Catalysis

Courtney A. Smith, Stephen E. Motika, Lukasz Wojtas, Xiaodong Shi *

<xmshi@usf.edu>

Department of Chemistry, University of South Florida, Tampa, Florida, 33620, USA.

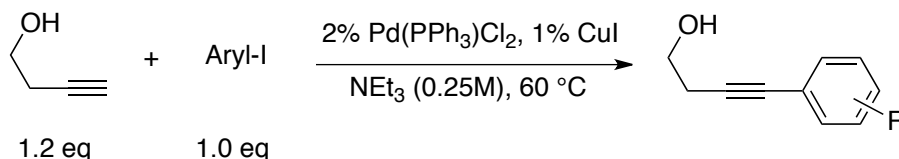
Table of Contents:

I. Materials and Methods	S2
II. ¹H-NMR Monitoring of Ring Expansion	S6
III. X-Ray Crystallographic Data for 6m, 7b, and 7g	S7
IV. Characterization Data	S14
V. NMR Spectral Data	S35

I. Materials and Methods

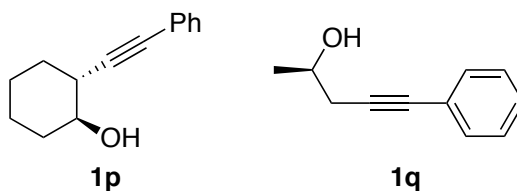
Unless otherwise noted, all reagents and solvents were obtained from a commercial provider and used without further purification. All reactions were run in oven-dried or flame-dried glassware under an atmosphere of N₂. Chloroform was dried over 3Å MS prior to use. ¹H- and ¹³C-NMR data was recorded on an Agilent 400 or 500 MHz NMR spectrometer. Reaction monitoring was carried out using either ¹H-NMR or analytical thin-layer chromatography. Analytical thin-layer chromatography was performed with pre-coated, glass-baked plates (250μ) and visualized by fluorescence or charring with potassium permanganate stain. Flash column chromatography was performed on 230-430 mesh silica gel. Chemical shifts for starting materials and products were recorded relative to internal TMS (0.00 ppm) for ¹H-NMR and CDCl₃ (77.00 ppm) for ¹³C-NMR data. HRMS were recorded on a LTQ-FTUHRA spectrometer.

1.1: General coupling procedure for synthesizing compounds **1a-1j**, **1l**.



Aryl-substituted homopropargyl alcohols (**1a-1j**; **1l**) were synthesized according to a modified literature procedure.¹ To a flame-dried flask was added aryl-iodide (1.0 eq, 4 mmol), alkynol (1.2 eq, 4.8 mmol), and triethylamine (0.25M). The mixture was placed under inert atmosphere and Pd(PPh₃)Cl₂ (2 mol %, 0.08 mmol) then CuI (1 mol %, 0.04 mmol) were sequentially added. The reaction was heated to 60 °C and stirred for 2-12 h, monitoring by TLC until completion. The crude was filtered through a short pad of silica, eluting with EtOAc (2 x 50 mL), then concentrated *in vacuo*. The mixture was purified via flash column chromatography on silica gel (eluting with 3:1 hexanes/ethyl acetate) to afford the corresponding aryl-substituted alkynols.

1.2: General procedure for synthesizing compounds **1p**, **1q**.

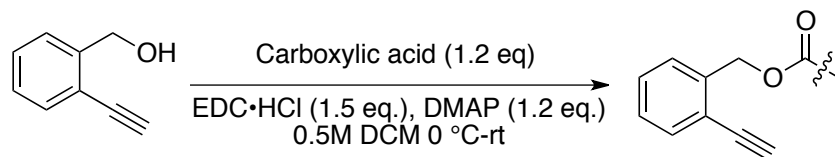


Chiral aryl-substituted homopropargyl alcohols (**1p**, **1q**) were synthesized according to literature procedure.²

¹ D. Y. Li, H. J. Chen, and P. N. Liu, *Angew. Chem.* 2016, **128**, 381.

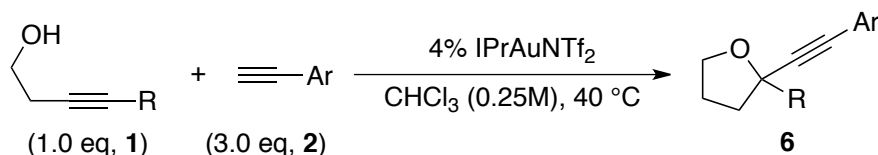
² R. D. Grigg, J. W. Rigoli, S. D. Pearce, and J. M. Schomaker, *Org. Lett.* 2012, **14**, 280.

1.3: General EDC coupling procedure for synthesizing compounds 2p, 2q, 2s, 2t.



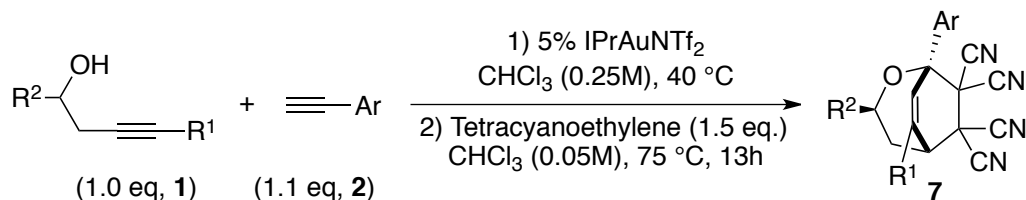
To an oven-dried flask was added the corresponding carboxylic acid (1.2 eq, 4.8 mmol) in dichloromethane (0.5M). DMAP (1.2 eq, 4.8 mmol) then alcohol (1.0 eq, 4.0 mmol) were added sequentially. The mixture was cooled to 0 °C. After cooling, EDC·HCl (1.5 eq, 6.0 mmol) was added slowly and the reaction was stirred at 0 °C for 5 minutes. The mixture was warmed to room temperature and stirred until completion, monitoring by TLC. The reaction was then diluted with water (20 mL), extracted with DCM (3 x 15 mL), and washed with brine (5 mL). The mixture was purified via flash column chromatography on silica gel (eluting with 5:1 hexanes/ethyl acetate) to afford the corresponding alkynes.

1.3: General procedure A for synthesizing compounds 6.



To a 2 mL vial charged with IPrAuNTf₂ (6.9 mg, 0.008 mmol) was sequentially added CHCl₃ (0.8 mL), **1** (1.0 eq, 0.2 mmol), and **2** (3.0 eq, 0.6 mmol). The reaction mixture was purged with argon for at least 1 minute and then stirred at 40 °C. After the reaction is complete (0.25h-7h), the mixture was directly purified via flash column chromatography on silica gel or by preparatory TLC to afford the desired product.

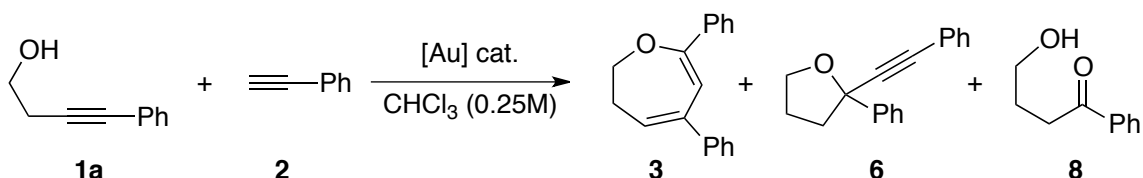
General procedure B for synthesizing compounds 7.



To a 2 mL vial charged with IPrAuNTf₂ (17.33 mg, 0.02 mmol) was sequentially added CHCl₃ (1.6 mL), **1** (1.0 eq, 0.4 mmol), and **2** (1.1 eq, 0.44 mmol). The reaction mixture was purged with argon for at least 1 minute and then stirred at 40 °C. After the reaction is complete (6h-16h), the mixture was passed through a short silica pad and collected in a

new 5mL-vial. The silica pad was rinsed with 2.0 mL CHCl₃. The dienophile was then added (0.6 mmol), the vial sealed securely with parafilm, and the resulting mixture was stirred for 13h at 75 °C. The reaction was concentrated *in vacuo* and purified via flash column chromatography on silica gel with 3:1 hexanes/ethyl acetate as eluent to afford the desired product.

Comprehensive screening table of catalytic conditions.

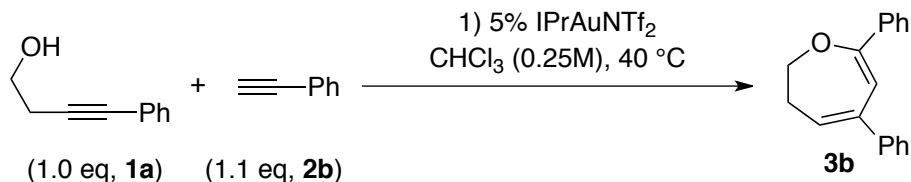


Entry	Conditions	T °C	t (h)	eq 2	convn (%)	yield (%) ^a		
						3	6	8
1	5% [XPhosAu(TA)]OTf, 1% Cu(OTf) ₂ , PhCH ₃	rt	24h	3	100	>5%	n.d.	n.d.
2	5% RuPhosAuNTf ₂	rt	12h	3	100	n.d.	47	n.d.
3	5% RuPhosAuNTf ₂	40	4h	3	100	n.d.	61	n.d.
4	5% IPrAuNTf ₂	40	4h	3	100	n.d.	91	n.d.
5	5% IPrAuNTf ₂	40	4h	2	100	56	36	n.d.
6	5% IPrAuNTTf ₂	40	11h	1.1	100	74	n.d.	n.d.
7	4% IPrAuNTf ₂	40	4h	3	100	n.d.	87	n.d.
8	1% IPrAuNTf ₂	40	4h	3	100	n.d.	34	53
9	5% PPh ₃ AuCl/AgOTf	40	4h	3	100	n.d.	32	61
10	5% PPh ₃ AuCl	40	4h	3	44	n.d.	n.d.	44
11	5% PPh ₃ AuNTf ₂	40	4h	3	100	n.d.	28	64
12	5% (tBu) ₃ PAuNTf ₂	40	4h	3	100	n.d.	32	60
13	(OAr) ₃ PAuNTf ₂ ^b	40	4h	3	100	n.d.	32	58
14	5% JohnPhosAuNTf ₂	40	4h	3	100	n.d.	27	66
15	5% XPhosAuNTf ₂	40	4h	3	100	n.d.	22	69
16	5% [RuPhosAu(TA-H)]OTf	40	4h	3	100	n.d.	29	47
17	5% [RuPhosAu(TA-Me)]OTf	40	4h	3	100	n.d.	37	32
18	5% AuCl ₃	40	4h	3	100	n.d.	n.d.	n.d.
19	10% AgSbF ₆	40	4h	3	46	n.d.	n.d.	23
20	10% AgOTf	40	4h	3	16	n.d.	n.d.	16

21	10% In(OTf) ₃	40	4h	3	100	n.d.	<5%	n.d.
22	10% HOTf	40	4h	3	100	n.d.	n.d.	15
23	10% ZnBr ₂	40	4h	3	45	n.d.	n.d.	38
24	5% CuBr	40	4h	3	0	n.d.	<5%	n.d.
25	5% CuI	40	4h	3	0	n.d.	<5%	n.d.
26	5% Cu(OTf) ₂	40	4h	3	0	n.d.	<5%	n.d.
27	5% Pd(OAc) ₂	40	4h	3	0	n.d.	<5%	n.d.

^a The yield was determined by ¹H-NMR using 1,3,5-trimethoxybenzene as the internal standard. ^b Ar = 2,4-Di-¹³C-ButylBenzene

II. $^1\text{H-NMR}$ Monitoring of Ring Expansion



To a 2 mL vial charged with IPrAuNTf₂ (17.33 mg, 0.02 mmol) was sequentially added CHCl₃ (1.6 mL), **1a** (1.0 eq, 0.4 mmol), and **2b** (1.1 eq, 0.44 mmol). The reaction mixture was purged with argon for at least 1 minute and then stirred at 40 °C. The yield of the products was determined via $^1\text{H-NMR}$ with 1,3,5-trimethoxybenzene as the internal standard.

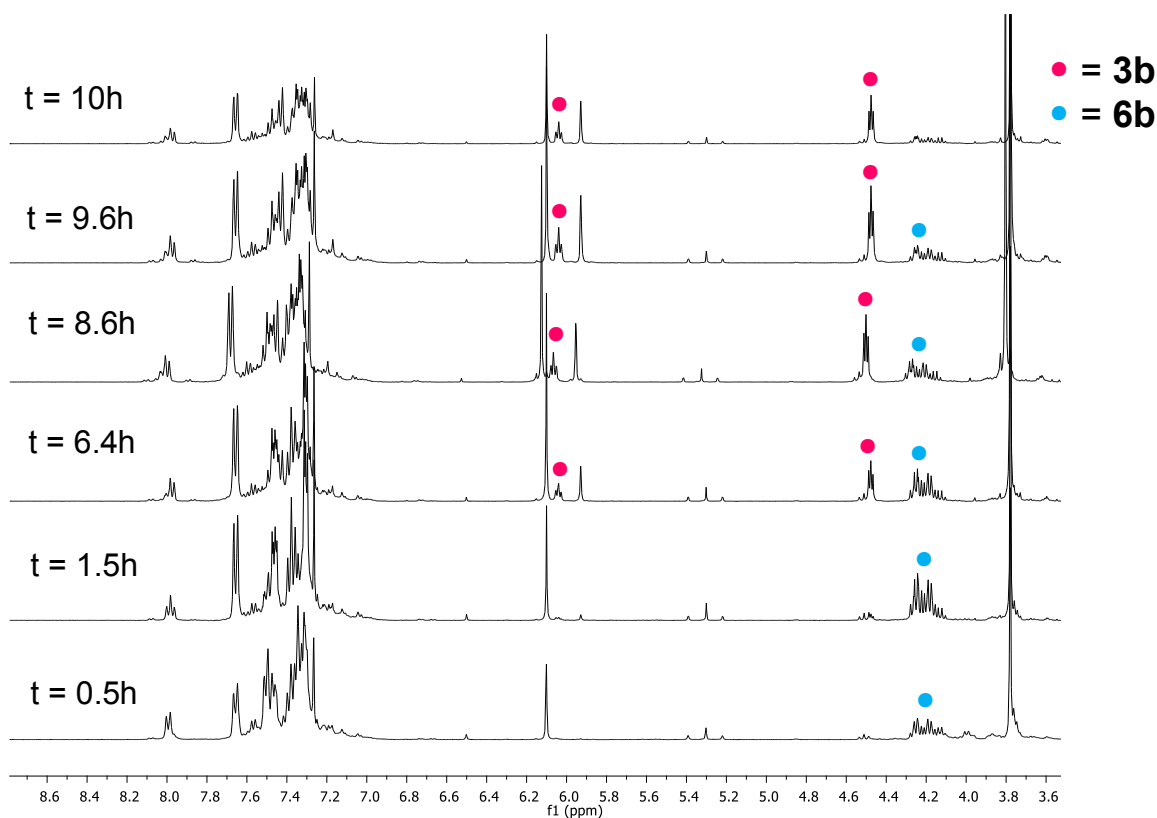


Figure 1. $^1\text{H-NMR}$ monitoring of ring expansion

III. X-Ray Crystallographic Data for 6m, 7b, and 7g

The X-ray diffraction data for all compounds were measured on Bruker D8 Venture PHOTON 100 CMOS system equipped with a Cu K α INCOATEC ImuS micro-focus source ($\lambda = 1.54178 \text{ \AA}$). Indexing was performed using *APEX3* [1] (Difference Vectors method). Data integration and reduction were performed using SaintPlus 6.01 [2]. Absorption correction was performed by a multi-scan method implemented in SADABS [3]. Space groups were determined using XPREP implemented in APEX3 [1]. Structures were solved using SHELXS-97 (direct methods) and refined using SHELXL-2014 [4-6] (full-matrix least-squares on F^2) through OLEX2 interface program [7]. All non-hydrogen atoms were refined anisotropically. Hydrogen atoms of -CH and -CH₂ groups were placed in geometrically calculated positions and were included in the refinement process using riding model with isotropic thermal parameters: Uiso(H) = 1.2Ueq(-CH,-CH₂).

Compound 7b: Crystal was a twin. The data have been integrated using two orientation matrices (twin law: -1 0 0 0 -1 0 0.4 0 1) and HKLF5 reflection file was used for refinement.

Compound 7g: The disordered -CF₃ group has been refined using distance (DFIX), angular (DANG) restraints and ADP restraints (RIGU). EADP constraint has been used for closely overlapping atoms C70A and C70B. Crystal data and refinement conditions are shown in Tables 1-3.

[1] Bruker (2016). *APEX3* (Version 2015.9). Bruker AXS Inc., Madison, Wisconsin, USA.

[2] Bruker (2016) SAINT V8.35A. Data Reduction Software.

[3] Sheldrick, G. M. (1996). *SADABS. Program for Empirical Absorption Correction*. University of Gottingen, Germany.

[4] Sheldrick, G.M. (1997) SHELXL-97. Program for the Refinement of Crystal

[5] Sheldrick, G.M. (1990) Acta Cryst. A46, 467-473

[6] Sheldrick, G. M. (2008). Acta Cryst. A64, 112-122.

[7] Dolomanov, O.V.; Bourhis, L.J.; Gildea, R.J.; Howard, J.A.K.; Puschmann, H., OLEX2: A complete structure solution, refinement and analysis program (2009). J. Appl. Cryst., 42, 339-341

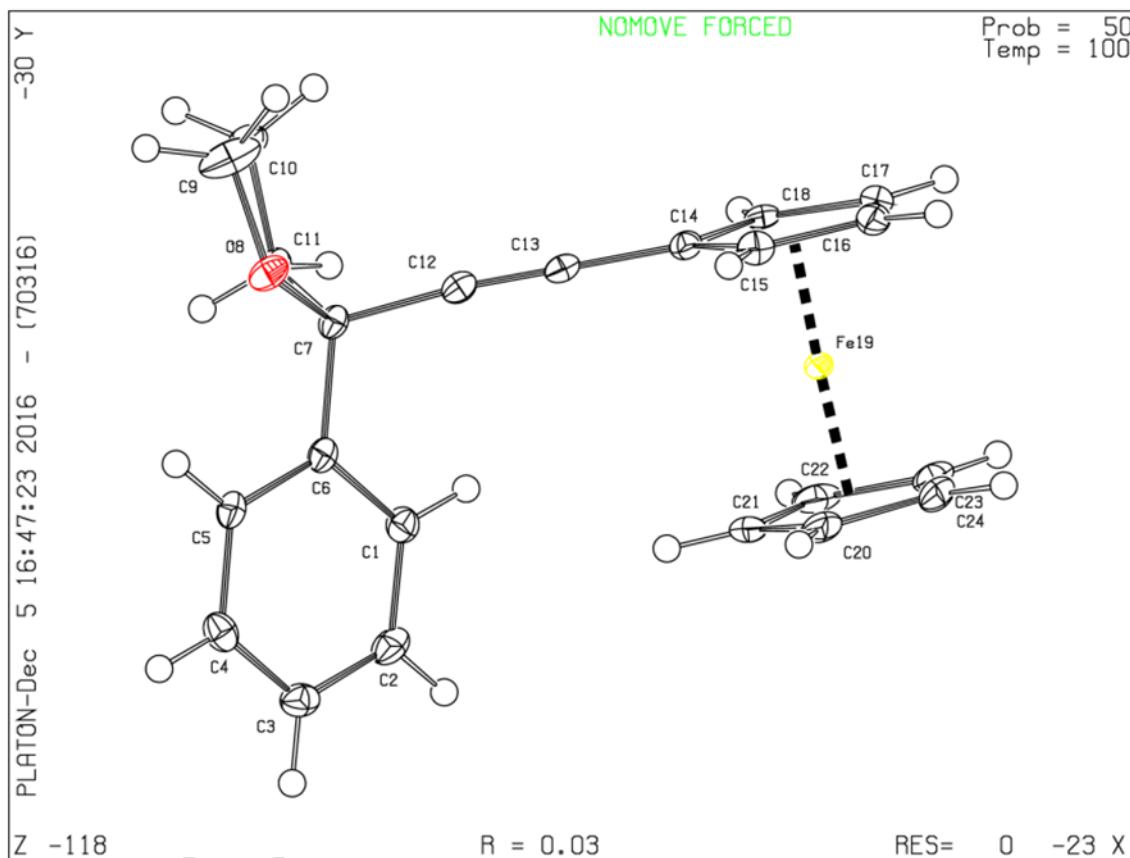


Figure 2. Perspective view of molecular structure $C_{22}H_{20}FeO$ with atom labeling.
CCDC 1521359

Table 1. Crystal data and structure refinement for 6m.

Identification code	6m
Empirical formula	C ₂₂ H ₂₀ FeO
Formula weight	356.23
Temperature/K	100.0
Crystal system	triclinic
Space group	P-1
a/Å	7.2823(2)
b/Å	10.5512(3)
c/Å	11.3059(3)
α/°	92.6290(10)
β/°	100.8210(10)
γ/°	101.1970(10)
Volume/Å ³	833.96(4)
Z	2
ρ _{calc} /g/cm ³	1.419
μ/mm ⁻¹	7.265
F(000)	372.0
Crystal size/mm ³	0.2 × 0.2 × 0.05
Radiation	CuKα (λ = 1.54178)
2Θ range for data collection/°	7.99 to 133.044
Index ranges	-8 ≤ h ≤ 8, -12 ≤ k ≤ 12, -12 ≤ l ≤ 13
Reflections collected	9825
Independent reflections	2911 [R _{int} = 0.0480, R _{sigma} = 0.0440]
Data/restraints/parameters	2911/0/217
Goodness-of-fit on F ²	1.033
Final R indexes [I ≥ 2σ (I)]	R ₁ = 0.0329, wR ₂ = 0.0734
Final R indexes [all data]	R ₁ = 0.0373, wR ₂ = 0.0761
Largest diff. peak/hole / e Å ⁻³	0.30/-0.32

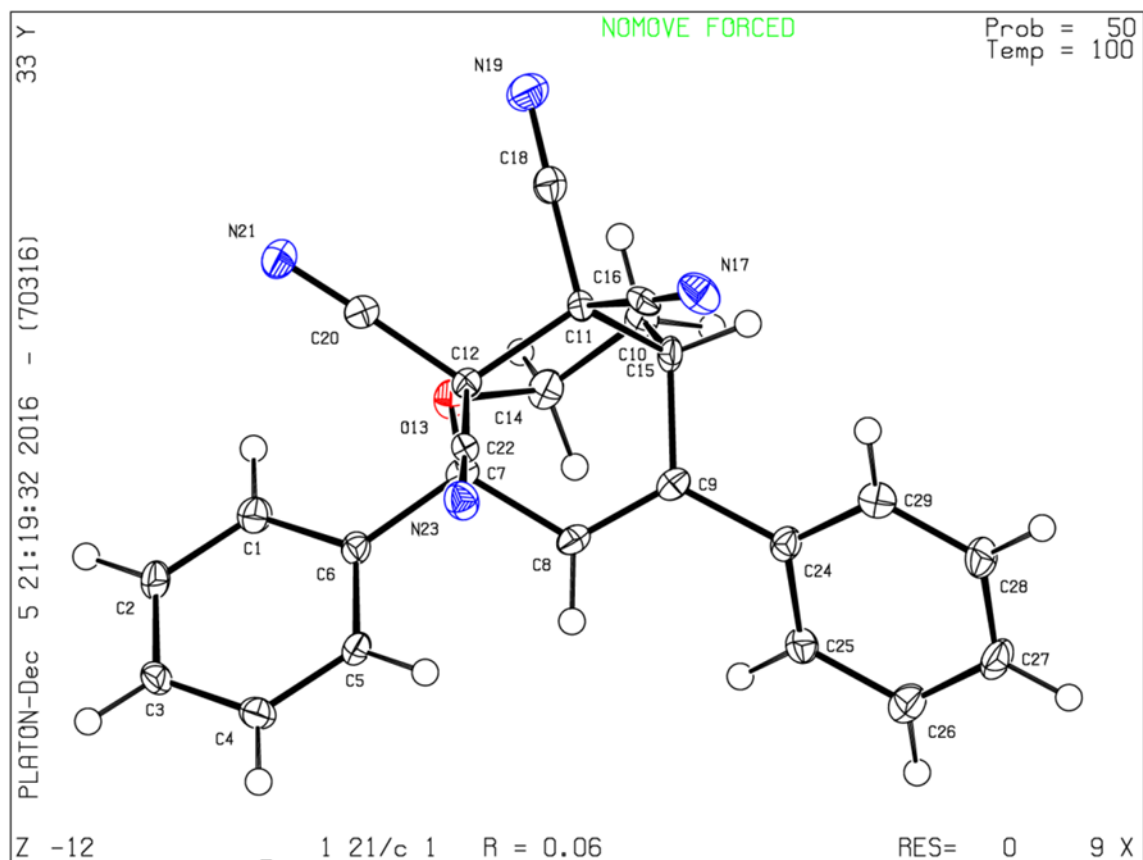


Figure 3. Perspective view of molecular structure $C_{24}H_{16}N_4O$ with atom labeling.
 CCDC 1521360

Table 2. Crystal data and structure refinement for 7b.

Identification code	7b
Empirical formula	C ₂₄ H ₁₆ N ₄ O
Formula weight	376.41
Temperature/K	100.03
Crystal system	monoclinic
Space group	P2 ₁ /c
a/Å	14.4429(4)
b/Å	8.4127(2)
c/Å	15.5871(4)
α/°	90
β/°	100.5300(10)
γ/°	90
Volume/Å ³	1862.00(8)
Z	4
ρ _{calc} /g/cm ³	1.343
μ/mm ⁻¹	0.680
F(000)	784.0
Crystal size/mm ³	0.276 × 0.192 × 0.088
Radiation	CuKα (λ = 1.54178)
2θ range for data collection/°	6.224 to 136.462
Index ranges	-17 ≤ h ≤ 17, 0 ≤ k ≤ 10, 0 ≤ l ≤ 18
Reflections collected	3385
Independent reflections	3385 [R _{int} = ?, R _{sigma} = 0.0493]
Data/restraints/parameters	3385/0/263
Goodness-of-fit on F ²	1.165
Final R indexes [I >= 2σ (I)]	R ₁ = 0.0649, wR ₂ = 0.1210
Final R indexes [all data]	R ₁ = 0.0819, wR ₂ = 0.1291
Largest diff. peak/hole / e Å ⁻³	0.26/-0.26

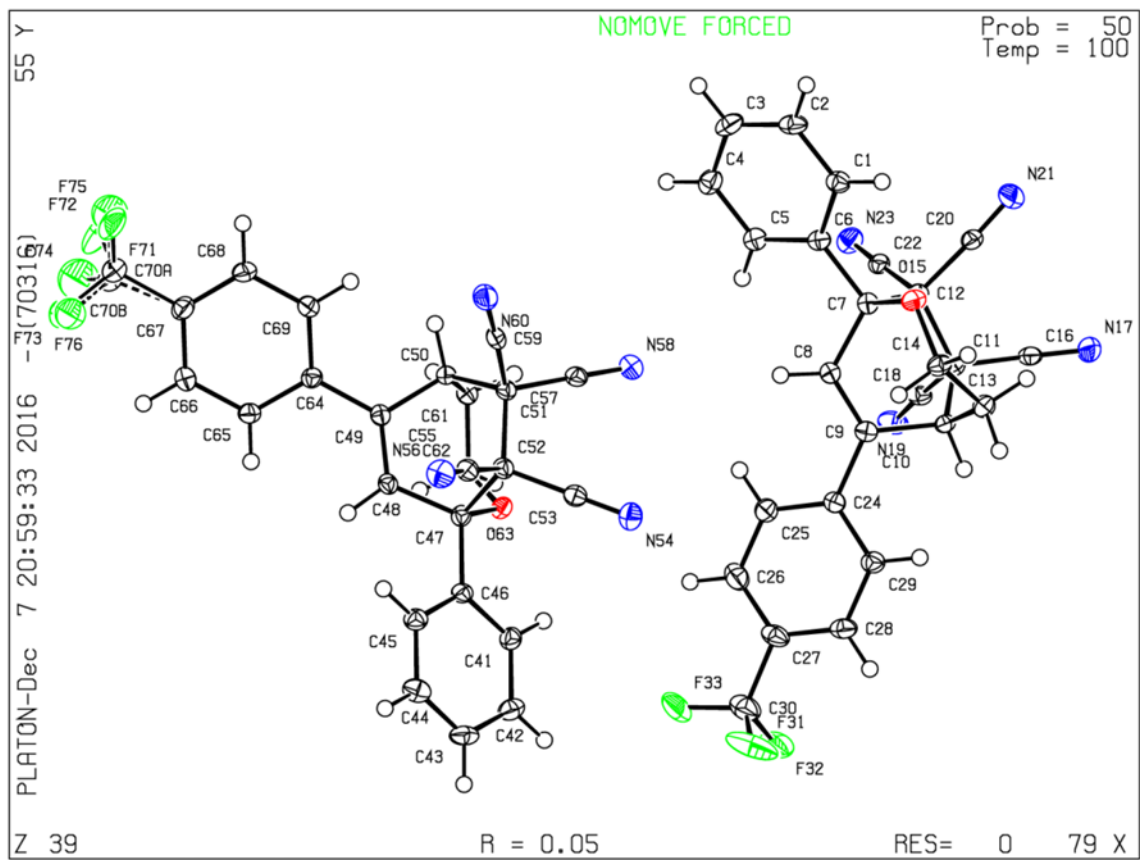
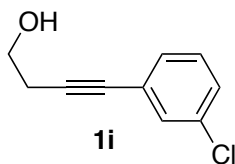


Figure 4. Perspective view of molecular structure $C_{25}H_{15}F_3N_4O$ with atom labeling.
CCDC 1521361

Table 3 - Crystal data and structure refinement for 7g.

Identification code	7g
Empirical formula	C ₂₅ H ₁₅ F ₃ N ₄ O
Formula weight	444.41
Temperature/K	100.04
Crystal system	triclinic
Space group	P-1
a/Å	10.3921(3)
b/Å	11.9914(3)
c/Å	17.5331(4)
α/°	80.3565(12)
β/°	89.9964(14)
γ/°	76.6551(13)
Volume/Å ³	2094.17(9)
Z	4
ρ _{calc} /g/cm ³	1.410
μ/mm ⁻¹	0.907
F(000)	912.0
Crystal size/mm ³	0.31 × 0.23 × 0.19
Radiation	CuKα (λ = 1.54178)
2θ range for data collection/°	5.116 to 154.388
Index ranges	-12 ≤ h ≤ 13, -15 ≤ k ≤ 15, -21 ≤ l ≤ 22
Reflections collected	30098
Independent reflections	8427 [R _{int} = 0.0517, R _{sigma} = 0.0460]
Data/restraints/parameters	8427/34/612
Goodness-of-fit on F ²	1.047
Final R indexes [I ≥ 2σ (I)]	R ₁ = 0.0463, wR ₂ = 0.0938
Final R indexes [all data]	R ₁ = 0.0719, wR ₂ = 0.1060
Largest diff. peak/hole / e Å ⁻³	0.31/-0.24

IV. Characterization Data

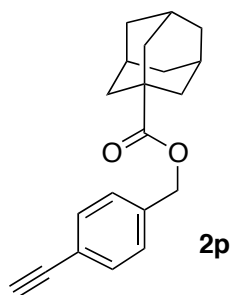


The title compound **1i** was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 3/1) as a pale-yellow oil (621 mg, 86% yield).

¹H NMR (500 MHz; CDCl₃): δ 7.45 – 7.34 (m, 1H), 7.28 – 7.12 (m, 3H), 3.83 – 3.73 (m, 2H), 3.01 (s, 1H), 2.68 – 2.60 (m, 2H).

¹³C NMR (126 MHz; CDCl₃): δ 133.97, 131.52, 129.76, 129.45, 128.14, 125.11, 88.09, 80.88, 60.93, 23.63.

HRMS calculated for C₁₀H₁₀ClO [M+H]⁺: 181.0420, Found: 181.0402

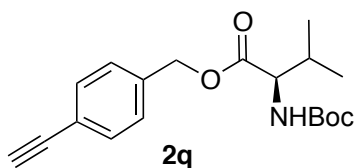


The title compound **2p** was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 5/1) as a white solid (754 mg, 64% yield).

¹H NMR (400 MHz; CDCl₃): δ 7.55 – 7.39 (m, 2H), 7.30 – 7.21 (m, 2H), 5.07 (s, 2H), 3.05 (s, 1H), 2.05 – 1.54 (m, 15H).

¹³C NMR (126 MHz; CDCl₃): δ 177.31, 137.29, 132.23, 127.47, 121.67, 83.31, 77.44, 65.17, 40.76, 38.80, 36.45, 27.89.

HRMS calculated for C₂₀H₂₃O₂ [M+H]⁺: 295.1698, Found: 295.1674

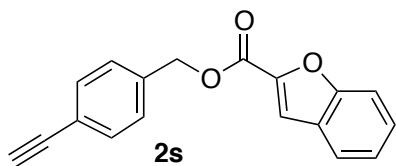


The title compound **2q** was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 5/1) as a white solid (716 mg, 54% yield).

¹H NMR (400 MHz; CDCl₃): δ 7.40 (d, *J* = 8.0 Hz, 2H), 7.24 (d, *J* = 8.0 Hz, 1.4H), 7.16 (d, *J* = 8.2 Hz, 0.6H), 5.16 – 4.97 (m, 3H), 4.25 – 4.16 (m, 1H), 3.05 (s, 1H), 2.24 – 1.96 (m, 1H), 1.37 (s, 9H), 0.87 (d, *J* = 6.8 Hz, 3H), 0.79 (d, *J* = 6.8 Hz, 3H).

¹³C NMR (126 MHz; CDCl₃): δ 172.14, 155.61, 136.11, 134.47, 132.22, 131.65, 129.95, 128.06, 122.35, 122.13, 83.13, 79.68, 77.86, 66.17, 58.55, 31.16, 28.26, 18.99, 17.50.

HRMS calculated for C₁₉H₂₆NO₄ [M+H]⁺: 332.1862, Found: 332.1846

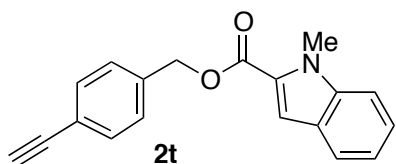


The title compound **2s** was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 5/1) as a white solid (906 mg, 82% yield).

¹H NMR (400 MHz; CDCl₃): δ 7.66 (d, *J* = 7.8 Hz, 1H), 7.59 – 7.47 (m, 4H), 7.47 – 7.36 (m, 3H), 7.29 (t, *J* = 7.5 Hz, 1H), 5.39 (s, 2H), 3.08 (s, 1H).

¹³C NMR (126 MHz; CDCl₃): δ 159.28, 155.81, 145.13, 136.04, 132.39, 128.20, 127.79, 126.86, 123.85, 122.86, 122.30, 114.39, 112.39, 83.17, 77.77, 66.42.

HRMS calculated for C₁₈H₁₂NaO₃ [M+Na]⁺: 299.0684, Found: 299.0670

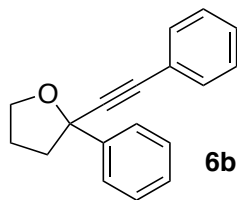


The title compound **2t** was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 5/1) as a white solid (706 mg, 61% yield).

¹H NMR (500 MHz; CDCl₃): δ 7.69 – 7.62 (m, *J* = 8.0 Hz, 1H), 7.50 (d, *J* = 8.2 Hz, 2H), 7.43 – 7.33 (m, 5H), 7.17 – 7.10 (m, 1H), 5.34 (s, 2H), 4.07 (s, 3H), 3.08 (s, 1H).

¹³C NMR (126 MHz; CDCl₃): δ 161.83, 139.78, 136.82, 132.35, 127.80, 127.32, 125.81, 125.19, 122.63, 121.99, 120.64, 110.64, 110.25, 83.265, 77.61, 65.56, 31.63.

HRMS calculated for C₁₉H₁₆NO₂ [M+H]⁺: 290.1181, Found: 290.1168



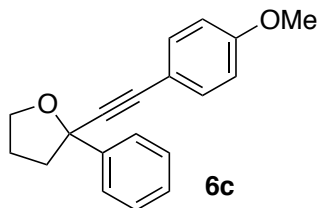
6b was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 24h, in CHCl₃.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 20/1) as a yellow oil (43 mg, 87%).

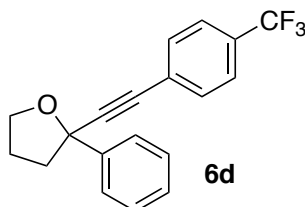
¹H NMR (500 MHz; CDCl₃): δ 7.70 – 7.63 (m, 2H), 7.50 – 7.44 (m, 2H), 7.40 – 7.33 (m, 2H), 7.32 – 7.27 (m, 4H), 4.32 – 4.10 (m, 2H), 2.62 (ddd, *J* = 12.2, 7.6, 4.7 Hz, 1H), 2.36 – 2.24 (m, 1H), 2.19 (dt, *J* = 12.0, 8.0 Hz, 1H), 2.13 – 2.02 (m, 1H).

¹³C NMR (126 MHz; CDCl₃): 143.64, 131.74, 128.25, 128.22, 128.18, 127.49, 125.32, 122.87, 91.26, 84.98, 80.92, 68.47, 43.02, 25.87.

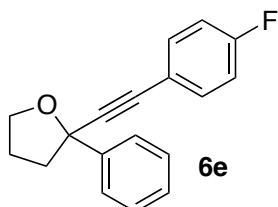
HRMS calculated for C₁₈H₁₇O [M+H]⁺: 249.1279, Found: 249.1278



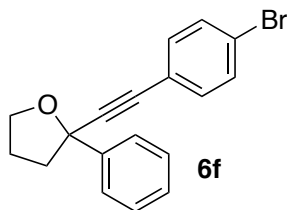
6c was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃. The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 20/1) as a pale-yellow oil (40 mg, 72%).
¹H NMR (500 MHz; CDCl₃): δ 7.64 (d, *J* = 7.8 Hz, 2H), 7.48 – 7.20 (m, 5H), 6.82 (d, *J* = 8.8 Hz, 2H), 4.20 (m, 2H), 3.81 (s, 3H), 2.65 – 2.49 (m, 1H), 2.33 – 1.98 (m, 3H).
¹³C NMR (126 MHz; CDCl₃): δ 159.53, 143.85, 133.17, 128.18, 127.40, 125.33, 114.98, 113.78, 89.81, 84.86, 80.95, 68.40, 55.26, 43.02, 25.86.
HRMS calculated for C₁₉H₁₉O₂ [M+H]⁺: 279.1385, Found: 279.1390



6d was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃. The title compound was isolated via preparatory TLC (Elution: hexanes/ethyl acetate = 20/1) as a yellow oil (54 mg, 86%).
¹H NMR: (500 MHz; CDCl₃): δ 7.67 – 7.60 (m, 2H), 7.58 – 7.52 (m, 4H), 7.38 (m, 2H), 7.36 – 7.28 (m, 1H), 4.22 (m, 2H), 2.62 (m, 1H), 2.32 – 2.18 (m, 2H), 2.12 – 2.04 (m, 1H).
¹³C NMR (126 MHz; CDCl₃): δ 143.26, 131.97, 130.00 (q, *J*_{C-F} = 32.58 Hz), 128.31, 127.64, 126.66, 125.19, 125.15, 122.79, 93.88, 83.54, 80.79, 68.62, 42.87, 25.86.
HRMS calculated for C₁₉H₁₆F₃O [M+H]⁺: 317.1153, Found: 317.1149



6e was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃. The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 20/1) as a yellow oil (44 mg, 82%).
¹H NMR: (500 MHz; CDCl₃): δ 7.67 – 7.60 (m, 2H), 7.47 – 7.40 (m, 2H), 7.37 (t, *J* = 7.6 Hz, 2H), 7.30 (t, *J* = 7.3 Hz, 1H), 7.03 – 6.95 (m, 2H), 4.25 – 4.15 (m, 2H), 2.60 (m, 1H), 2.31 – 2.13 (m, 2H), 2.11 – 2.01 (m, 1H).
¹³C NMR: (126 MHz; CDCl₃): δ 162.29 (d, *J*_{C-F} = 249.33 Hz), 143.59, 133.63 (d, *J*_{C-F} = 8.35 Hz), 128.24, 127.52, 125.25, 118.92 (d, *J*_{C-F} = 3.14 Hz), 115.45 (d, *J*_{C-F} = 22.04 Hz), 91.00, 83.85, 80.84, 68.50, 42.94, 25.86.
HRMS calculated for C₁₈H₁₆FO [M+H]⁺: 267.1185, Found: 267.1180.

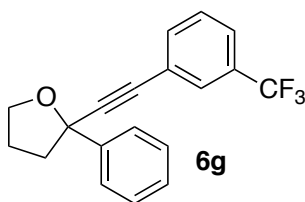


6f was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃. The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 30/1) as a yellow oil (44 mg, 68%).

¹H NMR (400 MHz; CDCl₃): δ 7.63 – 7.52 (m, 2H), 7.41 (d, *J* = 8.5 Hz, 2H), 7.35 (t, *J* = 7.5 Hz, 2H), 7.30 – 7.27 (m, 3H), 4.34 – 4.08 (m, 2H), 2.62 – 2.53 (m, 1H), 2.29 – 2.14 (m, 2H), 2.09 – 2.01 (m, 1H).

¹³C NMR (101 MHz; CDCl₃): δ 143.42, 133.16, 131.42, 128.24, 127.54, 125.20, 122.47, 121.79, 92.50, 83.82, 80.82, 68.52, 42.85, 25.83.

HRMS calculated for C₁₈H₁₆BrO [M+H]⁺: 327.0385, Found: 327.0371



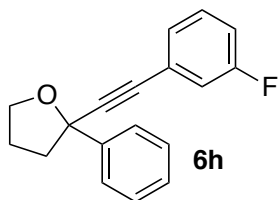
6g was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 20/1) as a colorless oil (39 mg, 61%).

¹H NMR (500 MHz; CDCl₃): δ 7.71 (s, 1H), 7.64 – 7.60 (m, 3H), 7.55 (d, *J* = 7.9 Hz, 1H), 7.45 – 7.36 (m, 3H), 7.31 (m, 1H), 4.21 (m, 2H), 2.62 (ddd, *J* = 12.2, 7.3, 4.7 Hz, 1H), 2.36 – 2.16 (m, 2H), 2.16 – 2.02 (m, 1H).

¹³C NMR (126 MHz; CDCl₃) δ 143.29, 134.83, 130.81 (q, *J*_{C-F} = 32.75 Hz), 128.72, 128.54, 128.31, 127.63, 125.19, 124.82, 123.77, 122.59, 93.01, 83.36, 80.77, 68.60, 42.86, 25.86

HRMS calculated for C₁₉H₁₆F₃O [M+H]⁺: 317.1153, Found: 317.1155



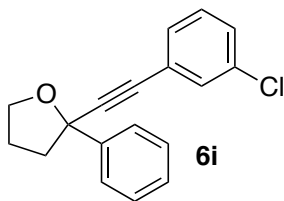
6h was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 20/1) as a colorless oil (47 mg, 89%).

¹H NMR (400 MHz; CDCl₃): δ 7.62 (d, *J* = 7.4 Hz, 2H), 7.37 (t, *J* = 7.5 Hz, 2H), 7.34 – 7.18 (m, 3H), 7.14 (d, *J* = 9.7 Hz, 1H), 7.07 – 6.95 (m, 1H), 4.32 – 4.10 (m, 2H), 2.66 – 2.54 (m, 1H), 2.35 – 1.97 (m, 3H).

¹³C NMR (126 MHz; CDCl₃): δ 163.24, 161.28, 143.38, 129.74 (d, *J*_{C-F} = 8.58 Hz), 128.27, 127.58, 125.23, 124.68 (d, *J*_{C-F} = 9.66 Hz) 118.54 (d, *J*_{C-F} = 22.82 Hz), 115.60 (d, *J*_{C-F} = 21.11 Hz), 92.28, 83.71, 80.80, 68.55, 42.91, 25.85.

HRMS calculated for C₁₈H₁₆FO [M+H]⁺: 267.1185, Found: 267.1189



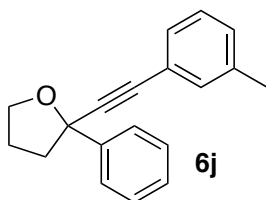
6i was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 1.5h, in CHCl₃.
The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 50/1) as a colorless oil (41 mg, 73%).

MW=282.77

¹H NMR (400 MHz; CDCl₃): δ 7.62 (d, *J* = 7.7 Hz, 2H), 7.44 (s, 1H), 7.40 – 7.15 (m, 6H), 4.41 – 4.02 (m, 2H), 2.73 – 2.43 (m, 1H), 2.48 – 1.92 (m, 3H).

¹³C NMR (101 MHz; CDCl₃): δ 143.33, 133.99, 131.59, 129.82, 129.39, 128.51, 128.25, 127.56, 125.20, 124.53, 92.57, 83.50, 80.78, 68.52, 42.88, 25.82.

HRMS calculated for C₁₈H₁₆ClO [M+H]⁺: 283.0890, Found: 283.0889

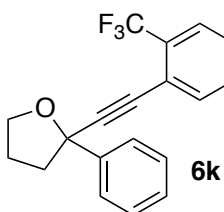


6j was synthesized with 4 mol% IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃.
The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 50/1) as a yellow oil (43 mg, 82%).

¹H NMR (500 MHz; CDCl₃): δ 7.66 – 7.61 (m, 2H), 7.36 (t, *J* = 7.6 Hz, 2H), 7.28 (t, *J* = 8.3 Hz, 1H), 7.18 (t, *J* = 7.6 Hz, 1H), 7.11 (d, *J* = 7.6 Hz, 1H), 4.27 – 4.13 (m, 2H), 2.59 (ddd, *J* = 12.2, 7.6, 4.6 Hz, 1H), 2.31 – 2.24 (m, 4H), 2.17 (m, 1H), 2.12 – 2.02 (m, 1H).

¹³C NMR (126 MHz; CDCl₃): δ 143.68, 137.85, 132.33, 129.12, 128.79, 128.20, 128.08, 127.46, 125.33, 122.66, 90.87, 85.16, 80.93, 68.43, 43.03, 25.87, 21.18.

HRMS calculated for C₁₉H₁₉O [M+H]⁺: 263.1436, Found: 263.1434

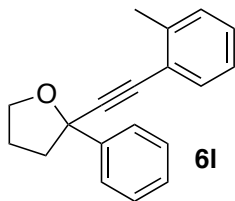


6k was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃.
The title compound was isolated via preparatory TLC (Elution: hexanes/ethyl acetate = 20/1) as a colorless oil (37 mg, 58%).

¹H NMR (500 MHz; CDCl₃): δ 7.67 – 7.61 (m, 3H), 7.57 (d, *J* = 7.7 Hz, 1H), 7.46 (t, *J* = 7.5 Hz, 1H), 7.38 (m, 3H), 7.29 (m, 1H), 4.28 – 4.14 (m, 2H), 2.64 (ddd, *J* = 11.3, 7.6, 3.7 Hz, 1H), 2.40 – 2.24 (m, 1H), 2.20 – 2.02 (m, 2H).

¹³C NMR (126 MHz; CDCl₃): δ 143.00, 133.96, 131.56 (q, *J*_{C-F} = 30.36 Hz), 131.33, 128.24, 128.02, 127.59, 125.72 (q, *J*_{C-F} = 4.97 Hz), 125.26, 124.65, 122.47, 121.14, 96.99, 81.00, 68.53, 42.94, 25.71.

HRMS calculated for C₁₉H₁₆F₃O [M+H]⁺: 317.1153, Found: 317.1148

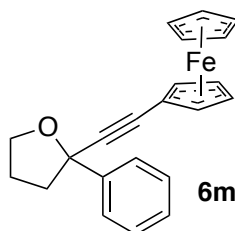


6l was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 15m, in CHCl₃. The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 50/1) as a pale-yellow oil (49 mg, 94%).

¹H NMR (500 MHz; CDCl₃): δ 7.67 (d, *J* = 7.5 Hz, 2H), 7.42 (d, *J* = 7.6 Hz, 1H), 7.38 (t, *J* = 7.6 Hz, 2H), 7.30 (t, *J* = 7.3 Hz, 1H), 7.23 – 7.17 (m, 2H), 7.15 – 7.09 (m, 1H), 4.29 – 4.16 (m, 2H), 2.61 (ddd, *J* = 12.3, 7.8, 4.5 Hz, 1H), 2.44 (s, 3H), 2.39 – 2.25 (m, 1H), 2.24 – 2.14 (m, 1H), 2.14 – 2.03 (m, 1H).

¹³C NMR (126 MHz; CDCl₃): δ 143.59, 140.23, 131.96, 129.33, 128.27, 128.20, 127.48, 125.44, 125.34, 122.63, 95.21, 84.07, 81.12, 68.35, 43.16, 25.87, 20.73.

HRMS calculated for C₁₉H₁₉O [M+H]⁺: 263.1436, Found: 263.1434.

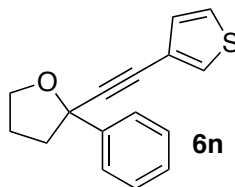


6m was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 5h, in CHCl₃. The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 20/1) as an orange solid (59 mg, 83%) and crystallized by slow evaporation of a DCM/Hexanes solution at room temperature.

¹H NMR (500 MHz; CDCl₃): δ 7.69 – 7.61 (m, 2H), 7.37 (t, *J* = 7.6 Hz, 2H), 7.29 (t, *J* = 7.3 Hz, 1H), 4.43 (t, *J* = 1.8 Hz, 2H), 4.27 – 4.13 (m, 9H), 2.55 (ddd, *J* = 12.1, 7.7, 4.6 Hz, 1H), 2.33 – 2.22 (m, 1H), 2.20 – 2.10 (m, 1H), 2.10 – 2.02 (m, 1H).

¹³C NMR (126 MHz; CDCl₃): δ 143.91, 128.15, 127.40, 125.36, 87.40, 83.64, 81.11, 71.57, 71.43, 69.80, 68.57, 68.26, 64.90, 43.04, 25.86.

HRMS calculated for C₂₂H₂₁FeO [M+H]⁺: 357.0942, Found: 357.0924

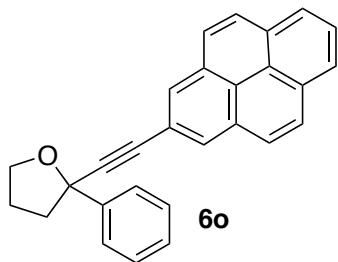


6n was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃. The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 20/1) as a yellow oil (47 mg, 93%).

¹H NMR (400 MHz; CDCl₃): δ 7.68 – 7.56 (m, 2H), 7.47 – 7.43 (m, 1H), 7.36 (t, *J* = 7.5 Hz, 2H), 7.30 – 7.23 (m, 2H), 7.11 (d, *J* = 5.0 Hz, 1H), 4.31 – 3.98 (m, 2H), 2.59 (ddd, *J* = 12.1, 7.2, 4.8 Hz, 1H), 2.43 – 1.93 (m, 3H).

¹³C NMR (101 MHz; CDCl₃): δ 143.62, 129.97, 128.77, 128.19, 127.45, 125.27, 125.09, 121.85, 90.81, 80.89, 80.03, 68.44, 42.89, 25.83.

HRMS calculated for C₁₆H₁₅OS [M+H]⁺: 255.0844, Found: 255.0840.



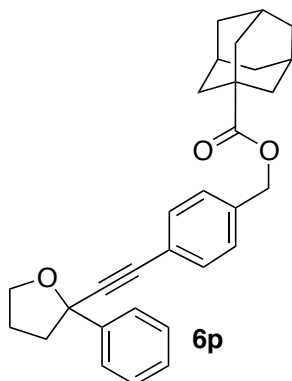
6o was synthesized with 4 mol% IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 20/1) as a pale-yellow oil (42 mg, 56%).

¹H NMR (400 MHz; CDCl₃): δ 8.49 (d, *J* = 9.1 Hz, 1H), 8.18 (t, *J* = 6.86, 2H), 8.14 – 7.97 (m, 7H), 7.82 – 7.74 (m, 1H), 7.42 (t, *J* = 7.6 Hz, 2H), 7.32 (t, *J* = 7.3 Hz, 1H), 4.38 (td, *J* = 8.1, 6.0 Hz, 1H), 4.29 – 4.24 (m, 1H), 2.79 (ddd, *J* = 12.2, 7.5, 4.6 Hz, 1H), 2.53 – 2.35 (m, 1H), 2.35 – 2.25 (m, 1H), 2.25 – 2.08 (m, 1H).

¹³C NMR (101 MHz; CDCl₃): δ 143.69, 132.01, 131.20, 131.18, 130.99, 129.76, 128.32, 128.08, 127.58, 127.19, 126.16, 125.53, 125.50, 125.41, 124.34, 124.24, 117.37, 96.97, 84.13, 81.34, 68.57, 43.22, 26.02.

HRMS calculated for C₂₈H₂₁O [M+H]⁺: 373.1592, Found: 373.1587



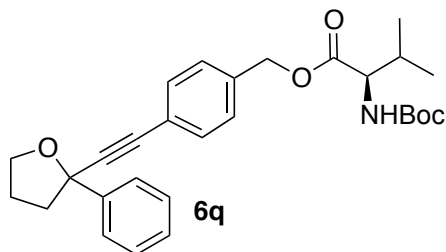
6p was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 1.5h, in CHCl₃.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 15/1) as a colorless oil (58 mg, 66%).

¹H NMR (500 MHz; CDCl₃): δ 7.67 – 7.60 (m, 2H), 7.44 (d, *J* = 8.2 Hz, 2H), 7.41 – 7.33 (m, 2H), 7.32 – 7.24 (m, 3H), 5.07 (s, 2H), 4.29 – 4.10 (m, 2H), 2.60 (ddd, *J* = 12.2, 7.5, 4.7 Hz, 1H), 2.28 (ddd, *J* = 11.7, 7.8, 6.1 Hz, 1H), 2.23 – 2.15 (m, 1H), 2.12 – 1.97 (m, 4H), 1.93 – 1.86 (m, 6H), 1.75 – 1.67 (m, 6H).

¹³C NMR (101 MHz; CDCl₃): δ 177.31, 143.56, 136.69, 131.83, 128.21, 127.47, 125.28, 122.48, 91.59, 84.64, 80.89, 68.46, 65.25, 42.98, 40.75, 38.79, 36.45, 27.89, 25.84.

HRMS calculated for C₃₀H₃₃O₃ [M+H]⁺: 441.2430, Found: 441.2412



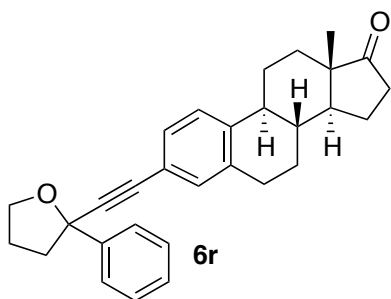
6q was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 7h, in CHCl₃.
The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 19/1) as a colorless oil (82 mg, 86%).

MW=477.60

¹H NMR (500 MHz; CDCl₃): δ 7.63 (d, *J* = 7.4 Hz, 2H), 7.44 (d, *J* = 8.1 Hz, 2H), 7.37 (t, *J* = 7.6 Hz, 2H), 7.33 – 7.27 (m, 3H), 5.19 (d, *J* = 12.5 Hz, 1H), 5.09 (d, *J* = 12.5 Hz, 1H), 4.99 (d, *J* = 8.8 Hz, 1H), 4.30 – 4.12 (m, 3H), 2.60 (ddd, *J* = 12.23, 7.46, 4.81 Hz, 1H), 2.36 – 2.00 (m, 4H), 1.44 (s, 9H), 0.93 (d, *J* = 6.8 Hz, 3H), 0.84 (d, *J* = 6.8 Hz, 3H).

¹³C NMR (126 MHz; CDCl₃): δ 172.21, 155.65, 143.54, 135.51, 131.91, 128.23, 128.09, 127.51, 125.27, 122.96, 91.88, 84.47, 80.87, 79.81, 68.51, 66.36, 58.53, 42.96, 31.25, 28.30, 25.86, 19.00, 17.49.

HRMS calculated for C₂₉H₃₆NO₅ [M+H]⁺: 478.2594, Found: 478.2582

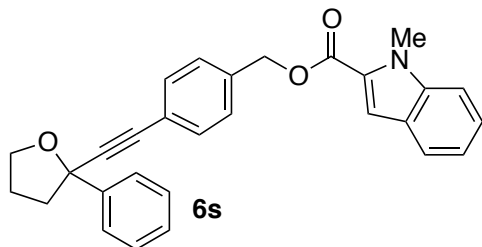


6r was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃.
The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 19/1) as a colorless oil (59 mg, 69%).

¹H NMR (500 MHz; CDCl₃): δ 7.69 – 7.61 (m, 2H), 7.40 – 7.34 (m, 2H), 7.32 – 7.19 (m, 4H), 4.27 – 4.13 (m, 2H), 2.87 (dd, *J* = 8.7, 3.9 Hz, 2H), 2.59 (ddd, *J* = 12.2, 7.6, 4.7 Hz, 1H), 2.51 (dd, *J* = 19.0, 8.7 Hz, 1H), 2.44 – 2.38 (m, 1H), 2.34 – 2.23 (m, 2H), 2.23 – 1.92 (m, 6H), 1.74 – 1.35 (m, 6H), 0.91 (s, 3H).

¹³C NMR (126 MHz; CDCl₃): δ 220.71, 143.71, 140.19, 136.45, 132.21, 129.03, 128.18, 127.43, 125.33, 125.23, 120.17, 90.57, 85.03, 80.95, 68.41, 50.48, 47.93, 44.41, 43.05, 37.96, 35.82, 31.53, 29.05, 26.32, 25.86, 25.58, 21.56, 13.82.

HRMS calculated for C₃₀H₃₃O₂ [M+H]⁺: 425.2481, Found: 425.2480



6s was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 1.5h, in CHCl₃.

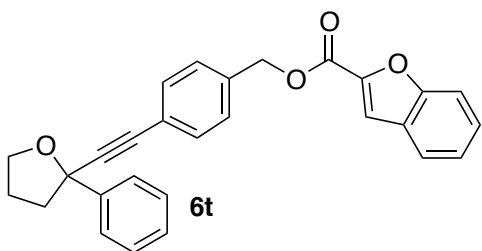
The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 15/1) as a white solid (69 mg, 79%).

MW= 435.52

¹H NMR (500 MHz; CDCl₃): δ 7.69 – 7.61 (m, 3H), 7.47 (d, *J* = 8.2 Hz, 2H), 7.42 – 7.33 (m, 7H), 7.30 – 7.27 (m, 1H), 7.16 – 7.13 (m, 1H), 5.34 (s, 2H), 4.30 – 4.12 (m, 2H), 4.07 (s, 3H), 2.61 (ddd, *J* = 12.2, 7.5, 4.8 Hz, 1H), 2.34 – 2.21 (m, 1H), 2.21 – 2.13 (m, 1H), 2.13 – 2.01 (m, 1H).

¹³C NMR (126 MHz; CDCl₃): δ 178.32, 161.85, 143.57, 139.77, 136.20, 131.97, 128.24, 127.79, 127.52, 125.82, 125.30, 125.15, 122.80, 122.62, 120.62, 110.61, 110.25, 91.78, 84.59, 80.91, 68.51, 65.68, 43.00, 31.63, 25.87.

HRMS calculated for C₂₉H₂₆NO₃ [M+H]⁺: 436.1913, Found: 436.1897



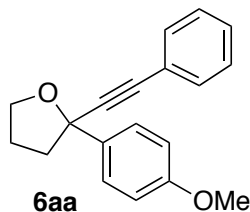
6t was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 6h, in CHCl₃.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 15/1) as a colorless oil (80 mg, 95%).

¹H NMR (500 MHz; CDCl₃): δ 7.68 (d, *J* = 7.8 Hz, 1H), 7.66 – 7.62 (m, 2H), 7.62 – 7.55 (m, 2H), 7.52 – 7.35 (m, 7H), 7.35 – 7.27 (m, 2H), 5.40 (s, 2H), 4.32 – 4.10 (m, 2H), 2.61 (ddd, *J* = 12.2, 7.5, 4.8 Hz, 1H), 2.36 – 1.97 (m, 3H).

¹³C NMR (126 MHz; CDCl₃): 159.30, 155.81, 145.19, 143.53, 135.43, 132.00, 128.24, 128.17, 127.76, 127.52, 126.87, 125.28, 123.83, 123.12, 122.85, 114.34, 112.39, 91.94, 84.49, 80.89, 68.51, 66.53, 42.98, 25.86.

HRMS calculated for C₂₈H₂₃O₄ [M+H]⁺: 423.1596, Found: 423.1586

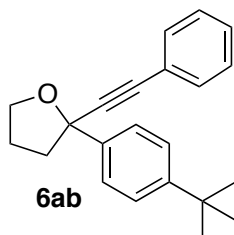


6aa was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃.
The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 20/1) as a yellow oil (37 mg, 67%).

¹H NMR (500 MHz; CDCl₃): δ 7.57 (d, *J* = 8.8 Hz, 2H), 7.47 – 7.43 (m, 2H), 7.33 – 7.26 (m, 3H), 6.90 (d, *J* = 8.8 Hz, 2H), 4.27 – 4.11 (m, 2H), 3.81 (s, 3H), 2.57 (ddd, *J* = 11.9, 7.5, 4.5 Hz, 1H), 2.37 – 2.01 (m, 3H).

¹³C NMR (126 MHz; CDCl₃): δ 158.99, 135.65, 131.72, 128.19, 128.17, 126.62, 122.90, 113.53, 91.34, 84.93, 80.65, 68.27, 55.29, 42.82, 25.85.

HRMS calculated for C₁₉H₁₉O₂ [M+H]⁺: 279.1385, Found: 279.1393



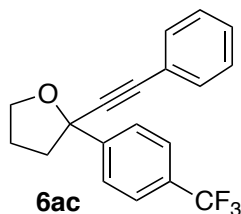
6ab was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 50/1) as a yellow oil (51 mg, 83%).

¹H NMR (500 MHz; CDCl₃): δ 7.57 (d, *J* = 8.5 Hz, 2H), 7.47 – 7.44 (m, 2H), 7.39 (d, *J* = 8.5 Hz, 2H), 7.31 – 7.28 (m, 3H), 4.26 – 4.13 (m, 2H), 2.59 (ddd, *J* = 12.2, 7.3, 4.6 Hz, 1H), 2.33 – 2.16 (m, 2H), 2.12 – 2.04 (m, 1H), 1.33 (s, 9H).

¹³C NMR (126 MHz; CDCl₃): δ 150.35, 140.50, 131.73, 128.15, 126.10, 125.13, 125.04, 122.97, 91.47, 84.82, 80.77, 77.25, 77.00, 76.74, 68.38, 42.74, 34.48, 31.35, 31.25, 25.90.

HRMS calculated for C₂₂H₂₅O [M+H]⁺: 305.1905, Found: 305.1903



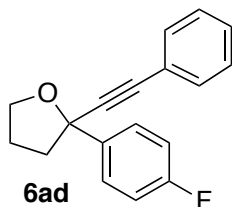
6ac was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 2.5h, in CHCl₃.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 20/1) as a yellow oil (47 mg, 74%).

¹H NMR (400 MHz; CDCl₃): δ 7.74 (d, *J* = 8.2 Hz, 2H), 7.61 (d, *J* = 8.3 Hz, 2H), 7.45 – 7.41 (m, 2H), 7.32 – 7.27 (m, 3H), 4.33 – 4.10 (m, 2H), 2.62 (ddd, *J* = 11.4, 7.5, 4.3 Hz, 1H), 2.37 – 2.20 (m, 1H), 2.20 – 1.99 (m, 2H).

¹³C NMR (101 MHz; CDCl₃): δ 147.75, 131.72, 129.69 (q, *J*_{C-F} = 32.26 Hz), 128.47, 128.22, 125.66, 125.48, 125.21 (q, *J*_{C-F} = 3.31 Hz), 122.44, 90.42, 85.41, 80.50, 68.65, 43.10, 25.81.

HRMS calculated for C₁₉H₁₆F₃O [M+H]⁺: 317.1153, Found: 317.1152

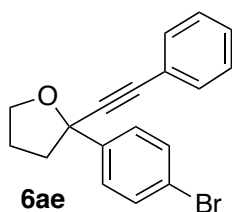


6ad was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃.
The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 50/1) as a pale-yellow oil (42 mg, 78%).

¹H NMR (400 MHz; CDCl₃): δ 7.64 – 7.55 (m, 2H), 7.47 – 7.39 (m, 2H), 7.31 – 7.26 (m, 3H), 7.03 (t, *J* = 8.7 Hz, 2H), 4.31 – 4.07 (m, 2H), 2.64 – 2.50 (m, 1H), 2.37 – 2.20 (m, 1H), 2.20 – 2.00 (m, 2H).

¹³C NMR (101 MHz; CDCl₃): δ 162.18 (d, *J*_{C-F} = 245.67 Hz), 139.39, 131.70, 128.33, 128.19, 127.09 (d, *J*_{C-F} = 8.07 Hz), 122.66, 114.95 (d, *J*_{C-F} = 21.47 Hz), 90.88, 85.18, 80.50, 68.40, 43.07, 25.82.

HRMS calculated for C₁₈H₁₆FO [M+H]⁺: 267.1185, Found: 267.1145



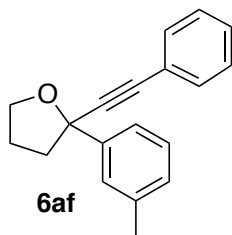
6ae was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 2.5h, in CHCl₃.
The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 20/1) as a pale-yellow oil (58 mg, 88%).

MW=327.22

¹H NMR (400 MHz; CDCl₃): δ 7.55 – 7.39 (m, 6H), 7.32 – 7.26 (m, 3H), 4.27 – 4.07 (m, 2H), 2.57 (ddd, *J* = 11.2, 7.5, 4.0 Hz, 1H), 2.33 – 2.18 (m, 1H), 2.18 – 1.97 (m, 2H).

¹³C NMR (101 MHz; CDCl₃): δ 142.82, 131.71, 131.27, 128.38, 128.20, 127.14, 122.57, 121.41, 90.61, 85.23, 80.47, 68.50, 43.00, 25.80.

HRMS calculated for C₁₈H₁₆BrO [M+H]⁺: 327.0385, Found: 327.0366

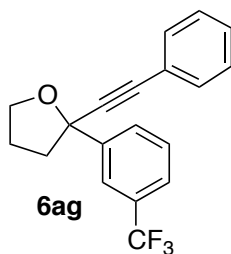


6af was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃.
The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 50/1) as a pale-yellow oil (40 mg, 76%).

¹H NMR (500 MHz; CDCl₃): δ 7.49 – 7.41 (m, 4H), 7.36 – 7.23 (m, 4H), 7.11 (d, *J* = 7.6 Hz, 1H), 4.27 – 4.14 (m, 2H), 2.60 (ddd, *J* = 12.2, 7.6, 4.6 Hz, 1H), 2.39 (s, 3H), 2.36 – 2.15 (m, 2H), 2.12 – 2.01 (m, 1H).

¹³C NMR (126 MHz; CDCl₃): δ 143.53, 137.85, 131.75, 128.24, 128.20, 128.16, 128.13, 125.89, 122.93, 122.44, 91.38, 84.84, 80.88, 68.39, 42.91, 25.84, 21.58.

HRMS calculated for C₁₉H₁₉O [M+H]⁺: 263.1436, Found: 263.1433

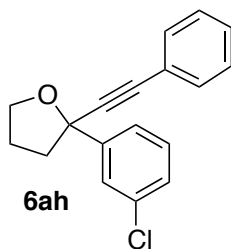


6ag was synthesized with 4 mol% IPrAuNTf₂, 3 equiv. alkyne, 60 °C, 3h, in CHCl₃. The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 20/1) as a yellow oil (51 mg, 81%).

¹H NMR (500 MHz; CDCl₃): δ 7.91 (s, 1H), 7.84 (d, *J* = 7.8 Hz, 1H), 7.56 (d, *J* = 7.8 Hz, 1H), 7.53 – 7.42 (m, 3H), 7.36 – 7.27 (m, 3H), 4.33 – 4.16 (m, 2H), 2.71 – 2.58 (m, 1H), 2.40 – 2.24 (m, 1H), 2.21 – 1.99 (m, 2H).

¹³C NMR (126 MHz; CDCl₃): δ 144.83, 131.75, 130.72 (q, *J*_{C-F} = 32.24 Hz), 128.88, 128.72, 128.49, 128.25, 124.38 (q, *J*_{C-F} = 3.41 Hz), 124.16 (q, *J*_{C-F} = 272.15 Hz), 122.47, 122.14 (q, *J*_{C-F} = 3.58 Hz), 90.40, 85.50, 80.51, 68.63, 43.16, 25.84.

HRMS calculated for C₁₉H₁₆F₃O [M+H]⁺: 317.1153, Found: 317.1151



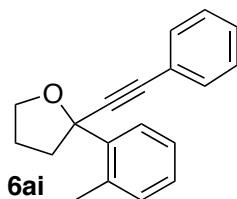
6ah was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 50/1) as a pale-yellow oil (48 mg, 84%).

¹H NMR (400 MHz; CDCl₃): δ 7.61 (s, 1H), 7.55 – 7.47 (m, 1H), 7.46 – 7.41 (m, 2H), 7.32 – 7.22 (m, 5H), 4.29 – 4.08 (m, 2H), 2.58 (ddd, *J* = 11.7, 7.5, 4.5 Hz, 1H), 2.35 – 2.19 (m, 1H), 2.19 – 1.96 (m, 2H).

¹³C NMR (101 MHz; CDCl₃): δ 145.82, 134.14, 131.73, 129.50, 128.39, 128.19, 127.62, 125.55, 123.58, 122.55, 90.54, 85.26, 80.39, 68.58, 43.08, 25.80.

HRMS calculated for C₁₈H₁₆ClO [M+H]⁺: 283.0890, Found: 283.0891



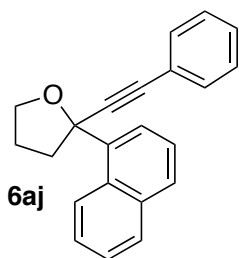
6ai was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 50/1) as a colorless oil (49 mg, 93%).

¹H NMR (400 MHz; CDCl₃): δ 7.67 (dd, *J* = 6.1, 2.9 Hz, 1H), 7.41 – 7.33 (m, 2H), 7.28 – 7.22 (m, 3H), 7.18 (d, *J* = 2.9 Hz, 3H), 4.30 – 4.17 (m, 1H), 4.13 – 3.97 (m, 1H), 2.79 – 2.69 (m, 1H), 2.61 (s, 3H), 2.30 – 2.15 (m, 2H), 2.07 – 1.94 (m, 1H).

¹³C NMR (101 MHz; CDCl₃): δ 141.45, 135.17, 131.70, 131.59, 128.11, 127.80, 127.40, 125.61, 124.51, 122.95, 91.09, 84.05, 79.78, 67.49, 40.36, 25.70, 21.08.

HRMS calculated for C₁₉H₁₉O [M+H]⁺: 263.1436, Found: 263.1438



6aj was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃.
The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 50/1) as a pale-yellow oil (48 mg, 81%).

¹H NMR (400 MHz; CDCl₃): δ 7.63 (d, *J* = 8.5 Hz, 1H), 7.86 (dd, *J* = 7.3, 3.2 Hz, 2H), 7.80 (d, *J* = 8.2 Hz, 1H), 7.58 – 7.42 (m, 3H), 7.40 – 7.32 (m, *J* = 6.6, 3.0 Hz, 2H), 7.26 – 7.19 (m, 3H), 4.37 – 4.09 (m, 2H), 3.00 – 2.87 (m, 1H), 2.56 – 2.42 (m, 1H), 2.42 – 2.20 (m, 1H), 2.08 – 1.93 (m, 1H).

¹³C NMR (101 MHz; CDCl₃): δ 138.97, 134.50, 131.64, 130.09, 128.76, 128.59, 128.13, 128.08, 126.10, 125.42, 125.29, 125.04, 122.88, 122.19, 91.95, 84.93, 79.66, 67.76, 41.31, 25.83.

HRMS calculated for C₂₂H₁₉O [M+H]⁺: 299.1436, Found: 299.1433

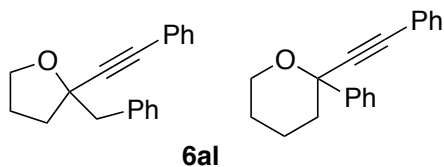


6ak was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 1.5h, in CHCl₃.
The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 20/1) as a colorless oil (36 mg, 91%).

¹H NMR (400 MHz; CDCl₃): δ 7.42 – 7.38 (m, 2H), 7.30 – 7.24 (m, 3H), 4.08 – 3.86 (m, 2H), 2.28 – 2.08 (m, 2H), 1.99 – 1.70 (m, 4H), 1.11 (t, *J* = 7.4 Hz, 3H).

¹³C NMR (101 MHz; CDCl₃): δ 131.48, 127.93, 127.79, 122.91, 91.17, 83.68, 80.41, 67.37, 38.17, 33.60, 25.30, 9.48.

HRMS calculated for C₁₄H₁₇O [M+H]⁺: 201.1279, Found: 201.1274

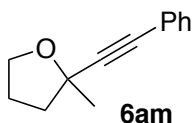


6al was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 1.5h, in CHCl₃.
The title compound was isolated as a mixture of isomers via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 33/1) as a yellow oil (46 mg, 87%, 10:3 *exo:endo* mixture).

¹H NMR (400 MHz; CDCl₃): 7.50 – 7.08 (m, 10H), 4.37 – 4.06 (m, 0.26H, *endo*), 4.06 – 3.80 (m, 1.74H), 3.18 – 3.03 (m, 1H), 2.24 – 1.78 (m, 5H).

¹³C NMR (126 MHz; CDCl₃): δ 149.63, 145.10, 143.48, 136.99, 135.86, 134.95, 131.58, 130.60, 129.45, 128.14, 128.06, 127.89, 127.17, 126.60, 126.15, 122.99, 121.18, 119.11, 93.44, 91.49, 84.80, 79.92, 69.76, 67.84, 46.50, 38.05, 33.88, 27.02, 25.51, 11.27.

HRMS calculated for C₁₉H₁₉O [M+H]⁺: 263.14359, Found: 263.1431

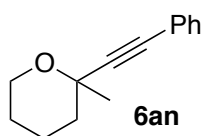


6am was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 2.5h, in CHCl₃. The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 25/1) as a colorless oil (31 mg, 84%).

¹H NMR (500 MHz; CDCl₃): δ 7.44 – 7.40 (m, 2H), 7.30 – 7.27 (m, 3H), 4.06 – 3.91 (m, 2H), 2.30 (ddd, *J* = 12.2, 8.2, 4.4 Hz, 1H), 2.22 – 2.09 (m, 1H), 2.05 – 1.95 (m, 1H), 1.92 – 1.81 (m, 1H), 1.64 (s, 3H).

¹³C NMR (126 MHz; CDCl₃): δ 131.66, 128.14, 128.04, 122.98, 92.33, 82.70, 76.39, 67.64, 40.16, 27.70, 25.70.

HRMS calculated for C₁₃H₁₅O [M+H]⁺: 187.1123, Found: 187.1117



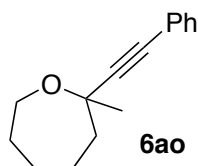
6an was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 30/1) as a pale-yellow oil (37 mg, 93%).

¹H NMR (500 MHz; CDCl₃): δ 7.48 – 7.42 (m, 2H), 7.33 – 7.28 (m, 3H), 4.02 (td, *J* = 11.5, 3.6 Hz, 1H), 3.85 – 3.78 (m, 1H), 1.96 – 1.83 (m, 2H), 1.76 – 1.69 (m, 1H), 1.61 – 1.53 (m, 6H).

¹³C NMR (126 MHz; CDCl₃): δ 131.70, 128.22, 128.13, 122.97, 90.28, 85.66, 70.71, 64.20, 38.09, 30.05, 25.47, 20.76.

HRMS calculated for C₁₄H₁₇O [M+H]⁺: 201.1279, Found: 201.1273



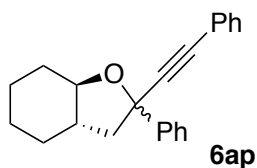
6ao was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 40/1) as a pale-yellow oil (34 mg, 79%).

¹H NMR (500 MHz; CDCl₃): δ 7.46 – 7.40 (m, 2H), 7.32 – 7.28 (m, 3H), 3.87 (ddd, *J* = 11.9, 8.6, 3.1 Hz, 1H), 3.75 (dt, *J* = 12.7, 4.0 Hz, 1H), 2.09 (dd, *J* = 14.7, 9.4 Hz, 1H), 1.96 (dd, *J* = 14.7, 8.2 Hz, 1H), 1.77 – 1.57 (m, 5H), 1.57 – 1.40 (m, 4H).

¹³C NMR (126 MHz; CDCl₃): δ 131.70, 128.17, 128.02, 123.09, 93.08, 83.07, 73.27, 64.88, 42.51, 30.86, 29.29, 28.47, 23.01.

HRMS calculated for C₁₅H₁₉O [M+H]⁺: 215.1436, Found: 215.1425



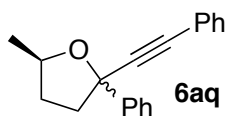
6ap was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃.

The title compound was isolated as a mixture of isomers via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 25/1) as a colorless oil (40 mg, 66%, 1:1 dr).

¹H NMR (500 MHz; CDCl₃): δ 7.69 – 7.60 (m, 2H), 7.50 – 7.43 (m, 2H), 7.37 (td, *J* = 7.7, 3.1 Hz, 2H), 7.32 – 7.27 (m, 4H), 3.64 (td, *J* = 10.7, 3.8 Hz, 0.5H), 3.51 – 3.39 (m, 0.5H), 2.77 (dd, *J* = 11.4, 5.2 Hz, 0.5H), 2.51 – 2.35 (m, 1H), 2.33 – 2.22 (m, 1H), 2.13 – 1.85 (m, 3.5H), 1.76 (dd, *J* = 6.4, 3.1 Hz, 1H), 1.63 – 1.50 (m, 1H), 1.44 – 1.08 (m, 3H).

¹³C NMR (126 MHz; CDCl₃): δ 145.65, 145.30, 131.77, 131.73, 128.32, 128.15, 128.11, 128.05, 127.25, 127.19, 125.48, 124.92, 123.04, 123.01, 93.33, 92.92, 85.17, 84.16, 84.00, 79.81, 78.99, 49.91, 49.76, 46.31, 44.25, 31.83, 31.12, 28.65, 25.70, 25.54, 24.33, 24.26.

HRMS calculated for C₂₂H₂₃O [M+H]⁺: 303.1749, Found: 303.1761



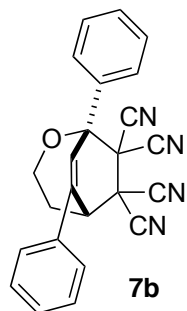
6aq was synthesized with 4 mol % IPrAuNTf₂, 3 equiv. alkyne, 40 °C, 4h, in CHCl₃.

The title compound was isolated as a mixture of isomers via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 30/1) as a colorless oil (50 mg, 95%, 3:2 dr).

¹H NMR (400 MHz; CDCl₃): δ 7.71 – 7.59 (m, 2H), 7.49 – 7.18 (m, 8H), 4.59 – 4.47 (m, 0.6H), 4.46 – 4.34 (m, 0.4H), 2.70 – 2.53 (m, 1H), 2.40 – 2.14 (m, 2H), 2.08 – 1.95 (m, 0.4H), 1.72 – 1.56 (m, 0.6H), 1.46 (d, *J* = 6.1 Hz, 1.2H), 1.40 (d, *J* = 6.1 Hz, 1.8H).

¹³C NMR (101 MHz; CDCl₃): δ 144.22, 143.96, 131.72, 131.62, 128.14, 127.39, 127.34, 125.37, 125.19, 123.05, 122.93, 92.56, 91.78, 84.80, 84.63, 81.04, 80.74, 75.70, 44.02, 43.27, 34.28, 32.89, 22.39, 21.04.

HRMS calculated for C₁₉H₁₉O [M+H]⁺: 263.1436, Found: 263.1430



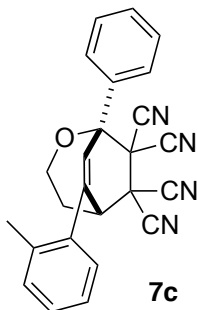
7b was synthesized using general procedure B.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 3/1) as a yellow oil (107 mg, 71% yield) and crystallized by slow evaporation of a CHCl₃/DCM/Hexanes solution at room temperature.

¹H NMR (400 MHz; CDCl₃): δ 7.83 – 7.76 (m, 2H), 7.61 – 7.45 (m, 8H), 6.75 (s, 1H), 4.19 – 4.10 (m, 2H), 3.80 (td, *J* = 12.3, 4.0 Hz, 1H), 2.93 – 2.81 (m, 1H), 2.08 – 1.97 (m, 1H).

¹³C NMR (126 MHz; CDCl₃): δ 144.83, 136.45, 135.11, 130.78, 130.36, 129.61, 128.72, 127.18, 125.89, 122.74, 112.92, 111.35, 111.22, 110.28, 81.89, 62.09, 47.12, 45.49, 29.70, 29.09.

HRMS calculated for C₂₄H₁₇N₄O [M+H]⁺: 377.1402, Found: 377.1394



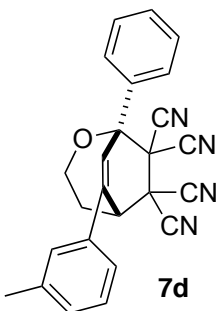
7c was synthesized using general procedure B.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 3/1) as a yellow oil (114 mg, 73% yield).

¹H NMR (400 MHz; CDCl₃): δ 7.79 – 7.71 (m, 2H), 7.56 – 7.47 (m, 3H), 7.39 – 7.19 (m, 4H), 6.56 (s, 1H), 4.24 (dd, *J* = 12.8, 6.8 Hz, 1H), 3.95 (td, *J* = 12.2, 4.3 Hz, 1H), 3.88 – 3.82 (m, 1H), 3.01 – 2.86 (m, 1H), 2.55 (s, 3H), 2.21 – 2.01 (m, 1H).

¹³C NMR (126 MHz; CDCl₃): δ 144.81, 136.32, 136.03, 135.40, 131.73, 130.36, 129.78, 128.77, 127.80, 127.07, 126.85, 126.72, 113.04, 111.55, 111.25, 110.22, 81.97, 62.50, 56.15, 48.52, 45.11, 29.60, 20.89.

HRMS calculated for C₂₅H₁₉N₄O [M+H]⁺: 391.1559, Found: 391.1563



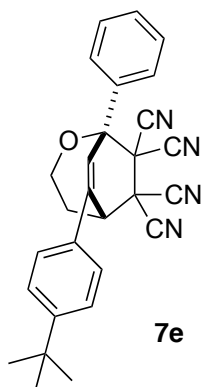
7d was synthesized using general procedure B.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 3/1) as a yellow oil (95 mg, 61% yield).

¹H NMR (500 MHz; CDCl₃): δ 7.83 – 7.71 (m, 2H), 7.57 – 7.45 (m, 3H), 7.45 – 7.27 (m, 4H), 6.72 (s, 1H), 4.18 – 4.07 (m, 2H), 3.79 (td, *J* = 12.3, 4.0 Hz, 1H), 2.95 – 2.80 (m, 1H), 2.42 (s, 3H), 2.08 – 1.99 (m, 1H).

¹³C NMR (126 MHz; CDCl₃): δ 144.89, 139.48, 136.47, 135.09, 131.54, 130.32, 129.46, 128.70, 127.17, 126.46, 122.96, 122.43, 112.87, 111.31, 111.20, 110.23, 81.86, 62.03, 47.13, 45.45, 35.85, 29.10, 21.47.

HRMS calculated for C₂₅H₁₉N₄O [M+H]⁺: 391.1559, Found: 391.1547



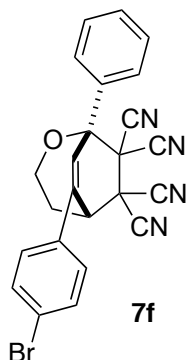
7e was synthesized using general procedure B.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 3/1) as a yellow oil (112 mg, 65% yield).

¹H NMR (400 MHz; CDCl₃): δ 7.89 – 7.68 (m, 2H), 7.68 – 7.44 (m, 7H), 6.72 (s, 1H), 4.28 – 4.01 (m, 2H), 3.90 – 3.64 (m, 1H), 2.99 – 2.73 (m, 1H), 2.15 – 1.89 (m, 1H), 1.45 – 1.27 (m, 9H).

¹³C NMR (101 MHz; CDCl₃): 154.43, 144.53, 136.54, 132.06, 130.28, 128.67, 127.16, 126.54, 125.58, 121.66, 112.89, 111.33, 111.24, 110.27, 81.91, 62.04, 61.99, 46.95, 45.46, 34.96, 34.92, 31.39, 31.10, 29.06.

HRMS calculated for C₂₈H₂₅N₄O [M+H]⁺: 433.2028, Found: 433.2012



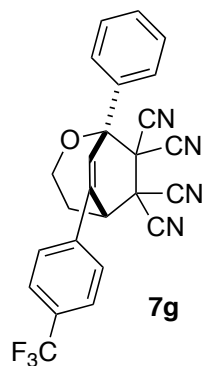
7f was synthesized using general procedure B.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 3/1) as a yellow solid (122 mg, 67% yield).

¹H NMR (500 MHz; CDCl₃): δ 7.81 – 7.77 (m, 2H), 7.65 (d, *J* = 8.3 Hz, 2H), 7.56 – 7.51 (m, 3H), 7.43 (d, *J* = 8.3 Hz, 2H), 6.78 (s, 1H), 4.22 – 4.08 (m, 2H), 3.78 (td, *J* = 12.3, 3.8 Hz, 1H), 2.95 – 2.78 (m, 1H), 2.08 – 1.99 (m, 1H).

¹³C NMR (126 MHz; CDCl₃): δ 143.83, 136.24, 133.94, 132.85, 130.45, 128.77, 127.40, 127.13, 125.32, 123.32, 112.87, 111.28, 111.07, 110.15, 81.85, 77.32, 77.07, 76.82, 62.09, 56.23, 46.98, 45.47, 29.03.

HRMS calculated for C₂₄H₁₆BrN₄O [M+H]⁺: 455.0508, Found: 455.0490



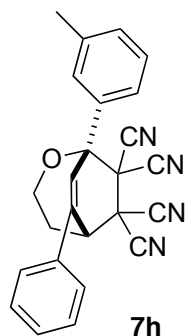
7g was synthesized using general procedure B.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 3/1) as a colorless oil (105 mg, 59% yield) and crystallized by slow evaporation of a CDCl_3 solution at room temperature.

$^1\text{H NMR}$ (500 MHz; CDCl_3): δ 7.82 – 7.73 (m, 4H), 7.66 (d, $J = 8.2$ Hz, 2H), 7.55 – 7.49 (m, 3H), 6.84 (s, 1H), 4.24 – 4.08 (m, 2H), 3.79 (td, $J = 12.3, 4.0$ Hz, 1H), 3.01 – 2.84 (m, 1H), 2.12 – 1.98 (m, 1H).

$^{13}\text{C NMR}$ (126 MHz; CDCl_3): δ 138.50, 136.01, 132.63 (q, $J_{\text{C-F}} = 33.08$ Hz), 130.54, 128.82, 127.89, 127.09, 126.66 (q, $J_{\text{C-F}} = 3.22$ Hz), 126.40, 125.17, 123.49 (q, $J_{\text{C-F}} = 272.64$ Hz), 112.83, 111.17, 110.92, 109.99, 81.77, 62.11, 56.23, 47.22, 45.46, 29.01.

HRMS calculated for $\text{C}_{25}\text{H}_{16}\text{F}_3\text{N}_4\text{O}$ $[\text{M}+\text{H}]^+$: 445.1276, Found: 445.1246



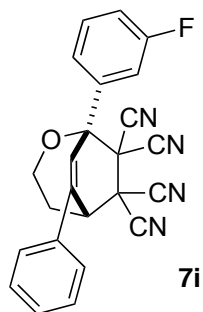
7h was synthesized using general procedure B.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 3/1) as a yellow solid (95 mg, 61% yield).

$^1\text{H NMR}$ (400 MHz; CDCl_3): 7.64 – 7.27 (m, 9H), 6.74 (s, 1H), 4.18 – 4.08 (m, 2H), 3.79 (td, $J = 12.2, 4.1$ Hz, 1H), 2.96 – 2.78 (m, 1H), 2.43 (s, 3H), 2.10 – 1.97 (m, 1H).

$^{13}\text{C NMR}$ (126 MHz; CDCl_3): δ 144.68, 138.60, 136.38, 135.16, 131.08, 130.73, 129.59, 128.58, 127.70, 125.88, 124.24, 122.91, 112.92, 111.33, 111.23, 110.29, 81.86, 62.02, 56.33, 47.15, 45.53, 29.05, 21.68.

HRMS calculated for $\text{C}_{25}\text{H}_{19}\text{N}_4\text{O}$ $[\text{M}+\text{H}]^+$: 391.1559, Found: 391.2515



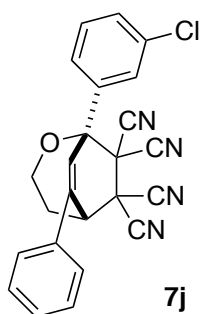
7i was synthesized using general procedure B.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 3/1) as a pale-yellow foam (107 mg, 68% yield).

¹H NMR (500 MHz; CDCl₃): δ 7.65 – 7.50 (m, 7H), 7.25 – 7.20 (m, 2H), 6.70 (d, *J* = 1.1 Hz, 1H), 4.30 – 4.09 (m, 2H), 3.82 (td, *J* = 12.3, 4.1 Hz, 1H), 3.01 – 2.60 (m, 1H), 2.16 – 1.99 (m, 1H).

¹³C NMR (126 MHz; CDCl₃): δ 162.65 (d, *J*_{C-F} = 248.16 Hz), 145.22, 138.95 (d, *J*_{C-F} = 7.09 Hz), 134.89, 130.93, 130.39 (d, *J*_{C-F} = 8.02 Hz), 129.65, 127.51 (d, *J*_{C-F} = 199.51 Hz), 125.88, 122.88 (d, *J*_{C-F} = 2.20 Hz), 122.13, 117.48 (d, *J*_{C-F} = 21.04 Hz), 114.84 (d, *J*_{C-F} = 24.33 Hz), 112.76, 111.11 (d, *J*_{C-F} = 17.34 Hz), 110.01, 81.47, 62.23, 56.07, 47.14, 45.40, 29.03.

HRMS calculated for C₂₄H₁₆FN₄O [M+H]⁺: 395.1308, Found: 395.1290



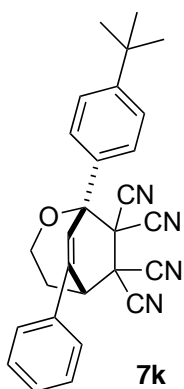
7j was synthesized using general procedure B.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 3/1) as a yellow oil (97 mg, 59% yield).

¹H NMR (500 MHz; CDCl₃): δ 7.81 (t, *J* = 1.7 Hz, 1H), 7.69 – 7.64 (m, 1H), 7.59 – 7.44 (m, 7H), 6.68 (s, 1H), 4.22 – 4.13 (m, 2H), 3.81 (td, *J* = 12.2, 4.1 Hz, 1H), 2.96 – 2.84 (m, 1H), 2.10 – 2.03 (m, 1H).

¹³C NMR (126 MHz; CDCl₃): δ 145.30, 138.40, 135.04, 134.88, 130.94, 130.61, 129.96, 129.64, 127.53, 125.87, 125.35, 122.01, 112.72, 111.13, 111.00, 109.93, 81.43, 62.21, 47.18, 45.38, 29.69, 29.01.

HRMS calculated for C₂₄H₁₆ClN₄O [M+H]⁺: 411.1013, Found: 411.0985



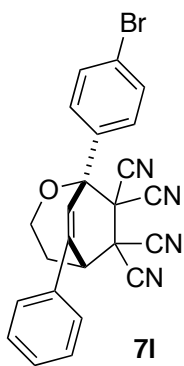
7k was synthesized using general procedure B.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 5/1) as a white solid (111 mg, 64% yield).

¹H NMR (500 MHz; CDCl₃): δ 7.80 – 7.69 (m, 2H), 7.61 – 7.48 (m, 7H), 6.78 (d, *J* = 1.1 Hz, 1H), 4.20 – 4.11 (m, 2H), 3.80 (td, *J* = 12.3, 4.0 Hz, 1H), 2.95 – 2.85 (m, 1H), 2.09 – 1.98 (m, 1H), 1.38 (s, 9H).

¹³C NMR (126 MHz; CDCl₃): δ 153.54, 144.55, 135.16, 133.47, 130.74, 129.60, 126.96, 125.89, 125.69, 123.07, 113.03, 111.44, 111.28, 110.41, 81.90, 62.03, 56.34, 47.17, 45.53, 34.80, 31.22, 29.16.

HRMS calculated for C₂₈H₂₅N₄O [M+H]⁺: 433.2028, Found: 433.2014



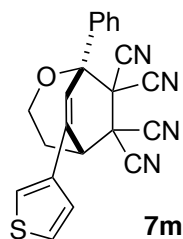
7l was synthesized using general procedure B.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 3/1) as a yellow solid (98 mg, 54% yield).

¹H NMR (500 MHz; CDCl₃): δ 7.65 (d, *J* = 9.2 Hz, 4H), 7.59 – 7.48 (m, 5H), 6.68 (s, 1H), 4.24 – 4.11 (m, 2H), 3.81 (td, *J* = 12.1, 4.0 Hz, 1H), 2.95 – 2.83 (m, 1H), 2.09 – 2.00 (m, 1H).

¹³C NMR (126 MHz; CDCl₃): δ 145.26, 135.43, 134.92, 131.94, 130.92, 129.64, 128.87, 125.86, 125.19, 122.04, 112.74, 111.22, 111.01, 110.03, 81.69, 62.16, 47.17, 45.31, 29.69, 29.01.

HRMS calculated for C₂₄H₁₆BrN₄O [M+H]⁺: 455.0508, Found: 455.0471



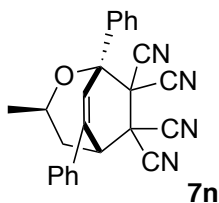
7m was synthesized using general procedure B.

The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 3/1) as a yellow solid (124 mg, 81% yield).

¹H NMR (500 MHz; CDCl₃): δ 7.71 (d, *J* = 2.0 Hz, 1H), 7.61 – 7.35 (m, 7H), 6.69 (s, 1H), 4.16 – 4.02 (m, 2H), 3.81 (td, *J* = 12.2, 4.0 Hz, 1H), 2.93 – 2.76 (m, 1H), 2.07 – 1.90 (m, 1H).

¹³C NMR (126 MHz; CDCl₃): δ 144.67, 138.45, 134.92, 130.84, 129.61, 127.13, 126.23, 125.91, 122.89, 112.90, 111.39, 111.13, 110.35, 81.30, 62.12, 55.87, 47.12, 45.34, 29.14.

HRMS calculated for C₂₂H₁₅N₄OS [M+H]⁺: 383.0967, Found: 383.0951



7n was synthesized using general procedure B.

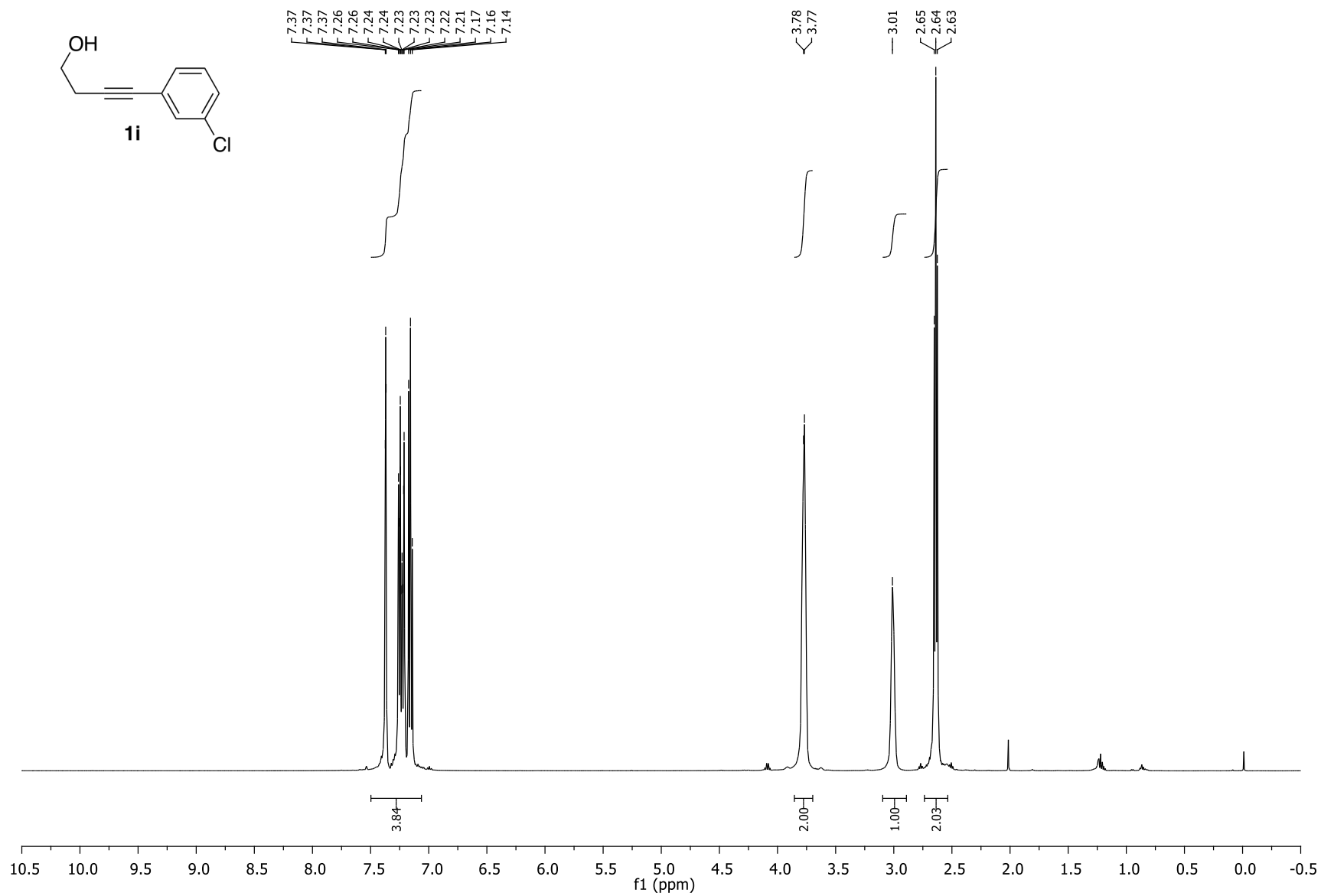
The title compound was isolated via flash chromatography on silica gel (Elution: hexanes/ethyl acetate = 3/1) as a pale-yellow oil (116 mg, 74% yield).

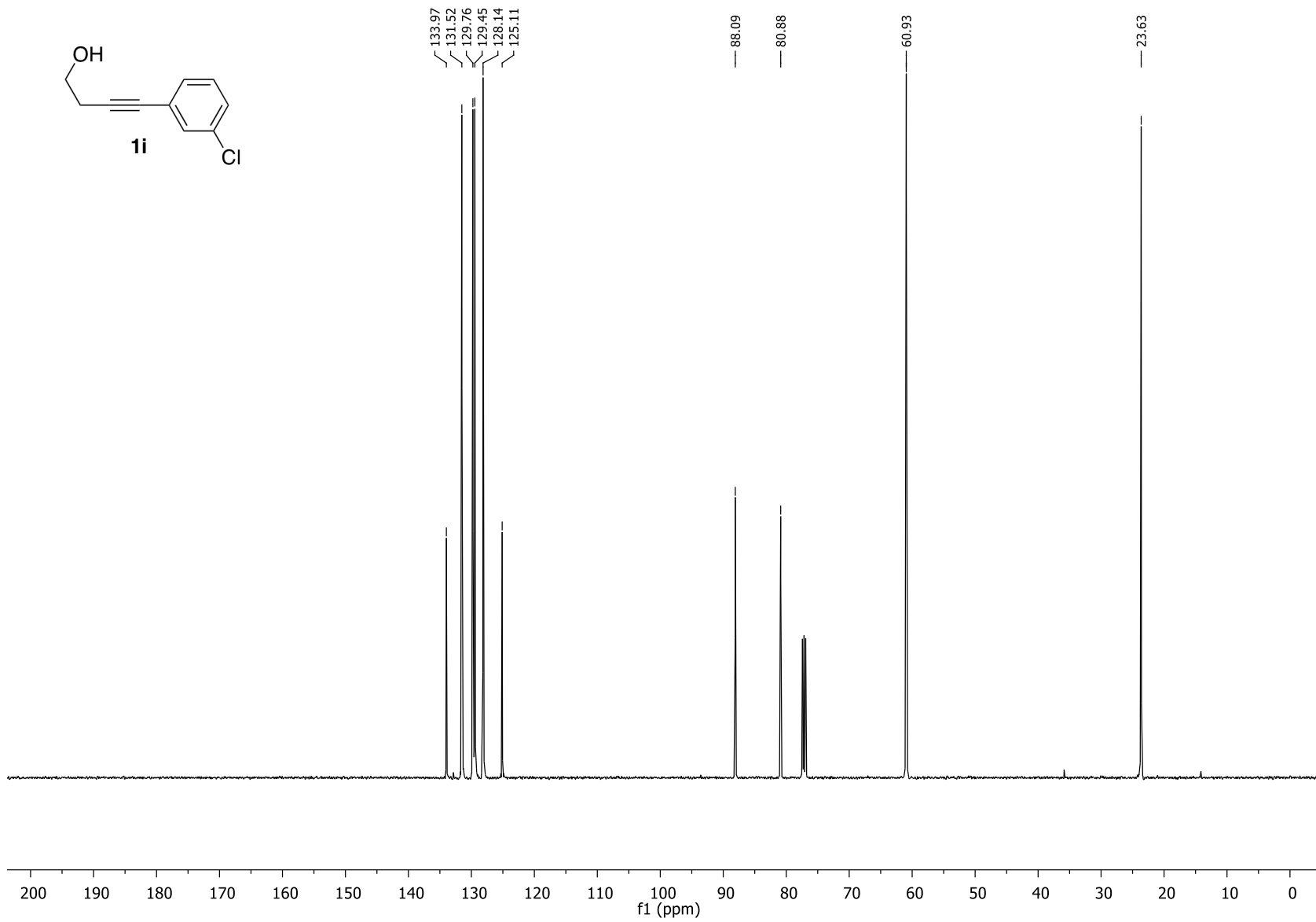
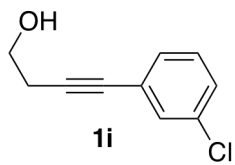
¹H NMR (500 MHz; CDCl₃): δ 7.90 – 7.77 (m, 2H), 7.63 – 7.43 (m, 8H), 6.78 (s, 1H), 4.11 (d, *J* = 6.4 Hz, 1H), 4.06 – 3.97 (m, 1H), 2.55 (dd, *J* = 15.4, 10.7 Hz, 1H), 2.27 – 2.10 (m, 1H), 1.43 (d, *J* = 6.2 Hz, 3H).

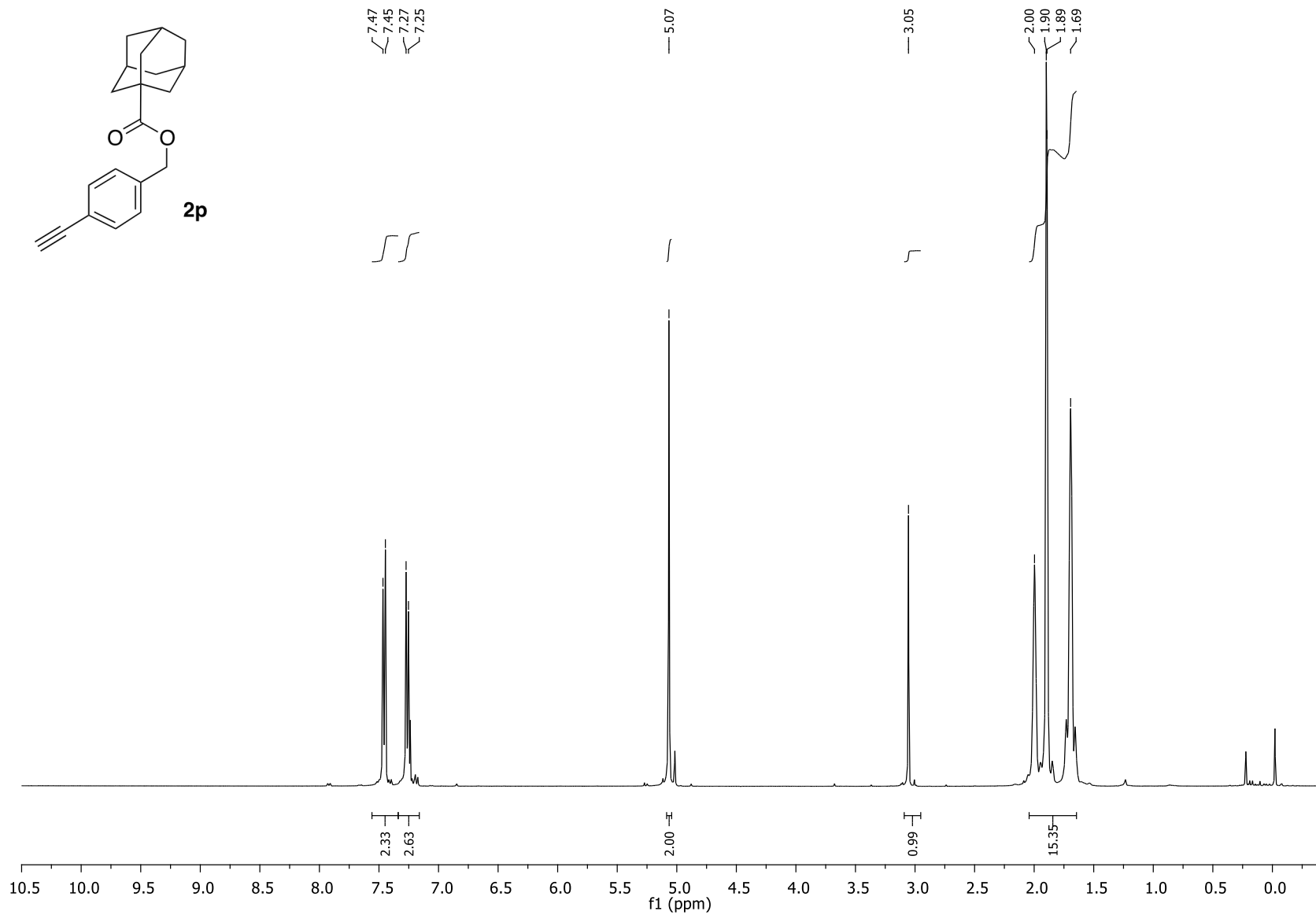
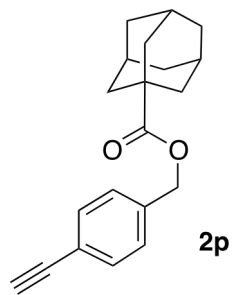
¹³C NMR (126 MHz; CDCl₃): δ 145.03, 136.70, 135.11, 130.70, 130.30, 129.58, 128.75, 127.14, 125.85, 123.55, 112.95, 111.39, 111.21, 110.26, 81.39, 77.30, 77.04, 76.79, 69.75, 56.34, 46.76, 45.32, 36.37, 20.27.

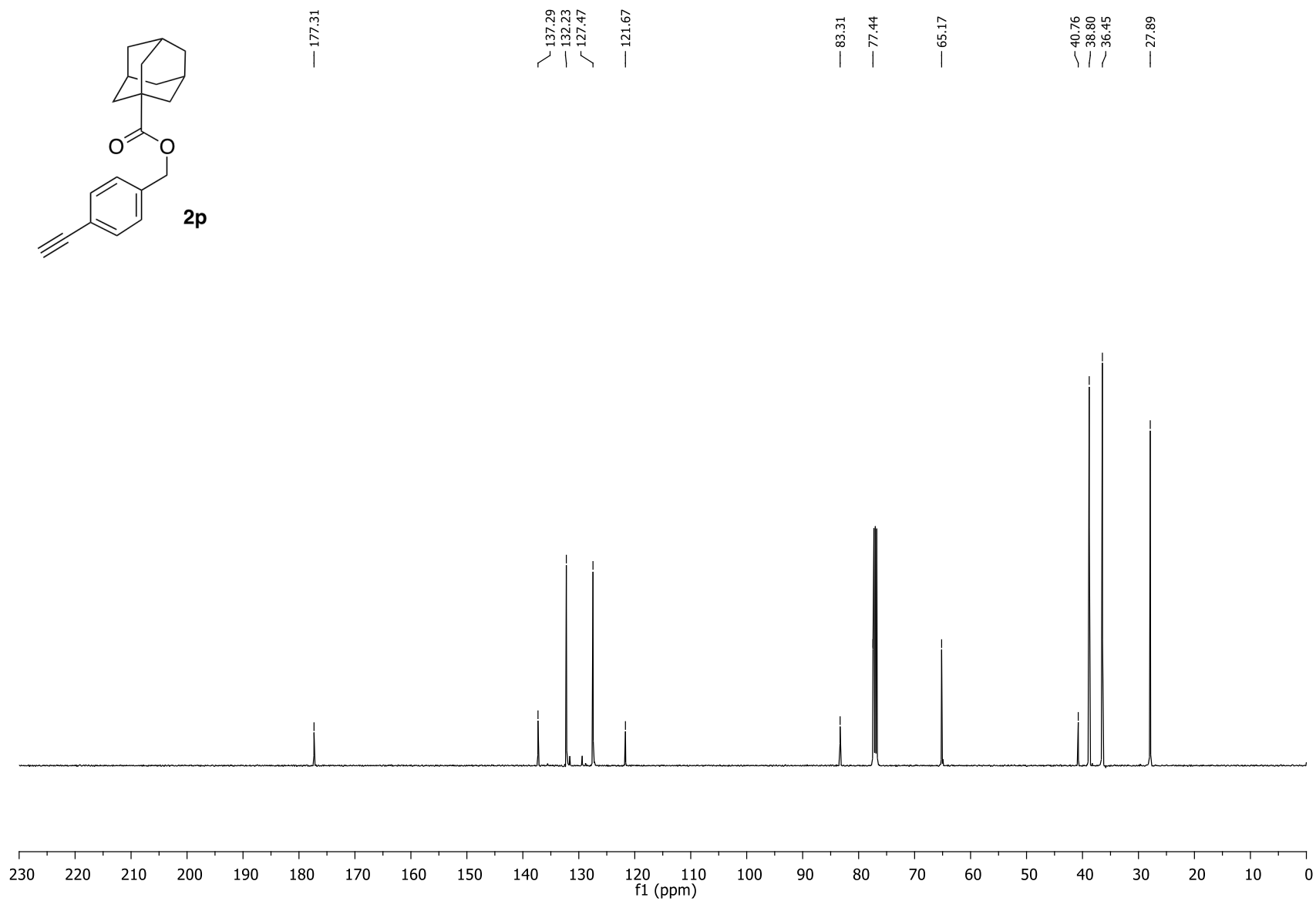
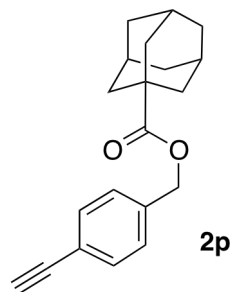
HRMS calculated for C₂₅H₁₉N₄O [M+H]⁺: 391.1559, Found: 391.1542

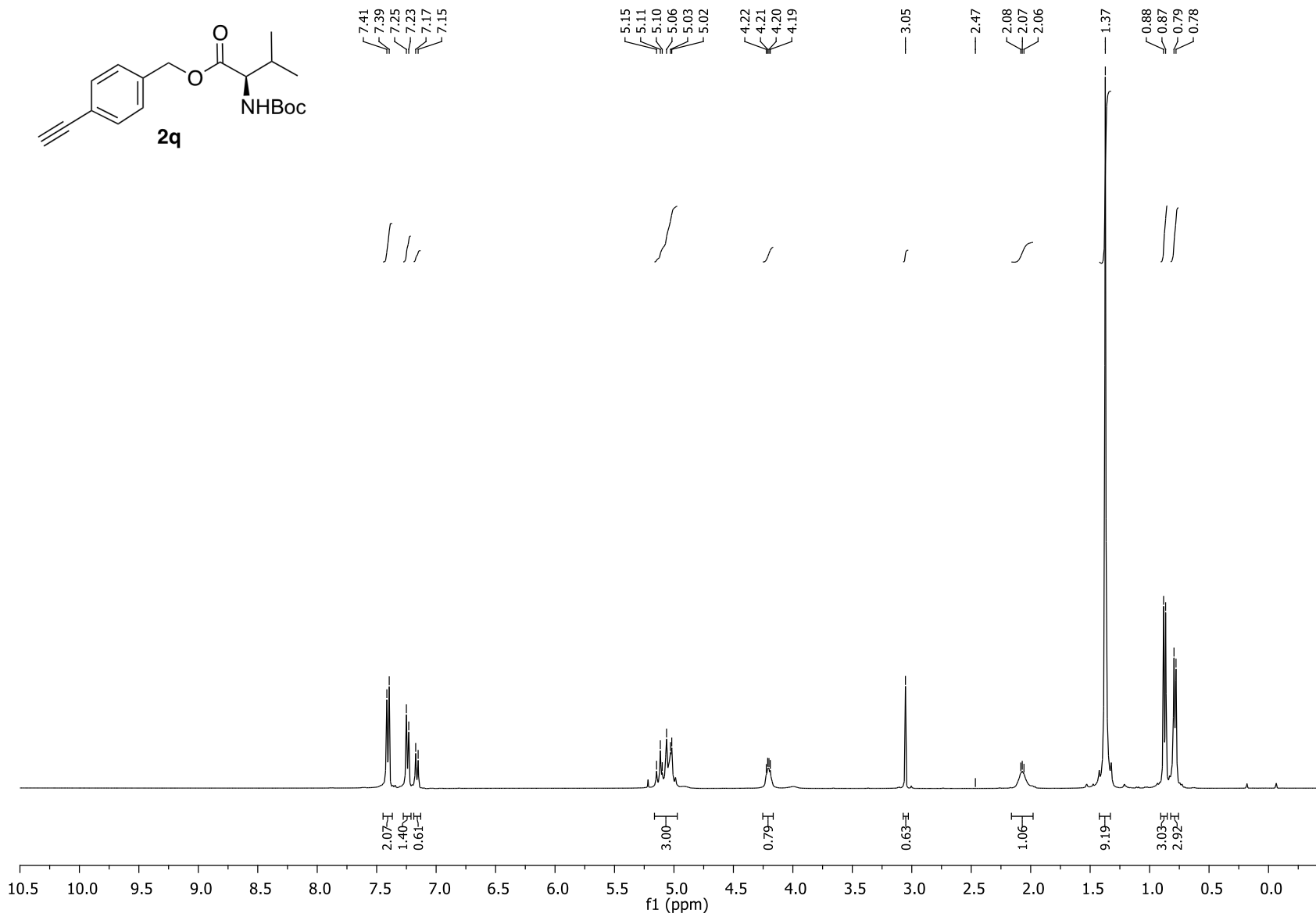
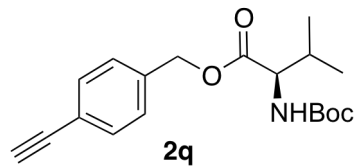
V. NMR Spectral Data

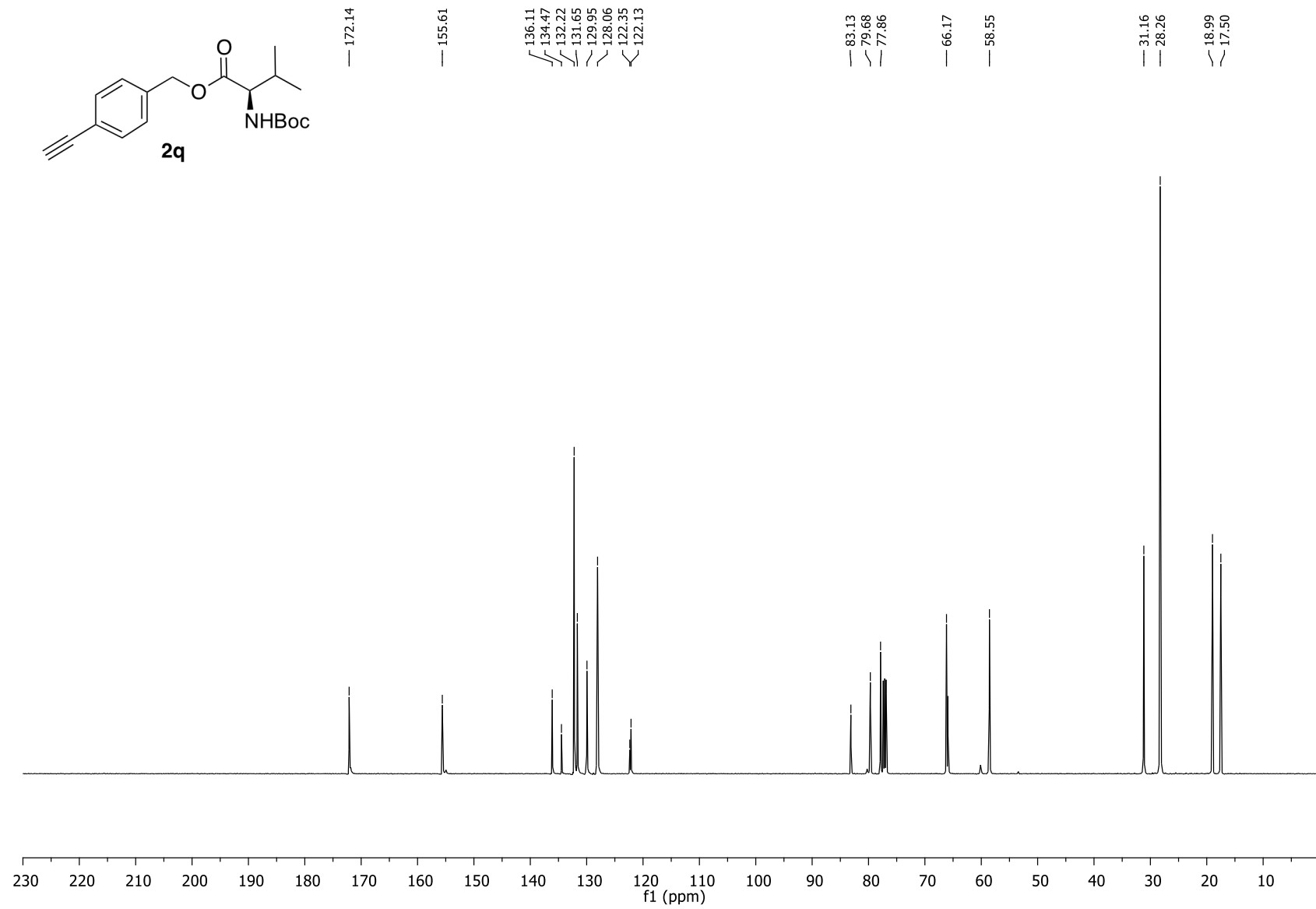
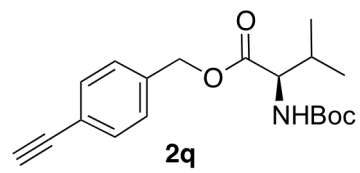


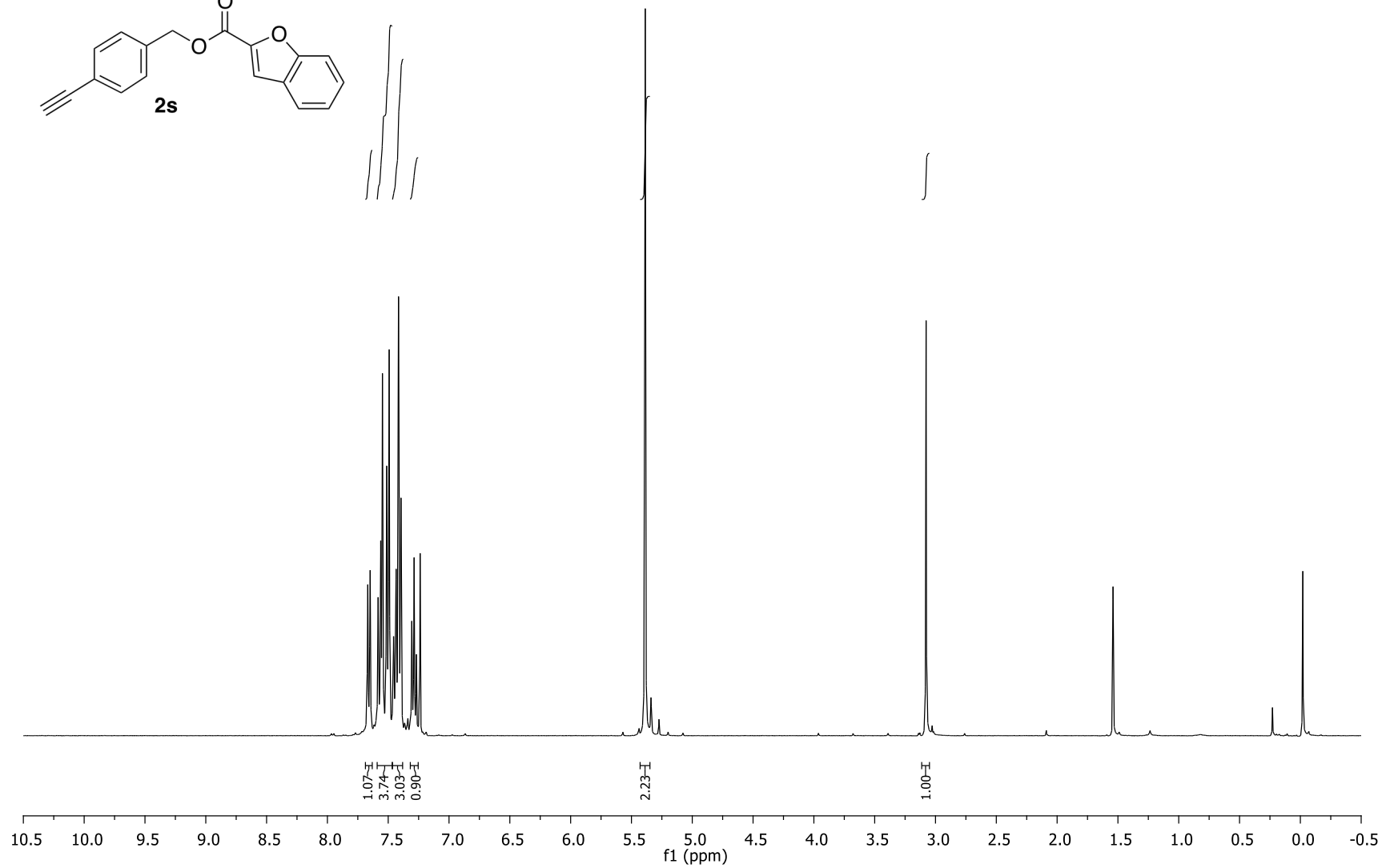
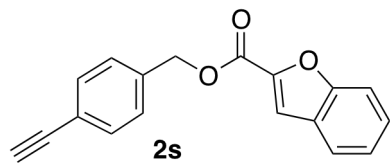


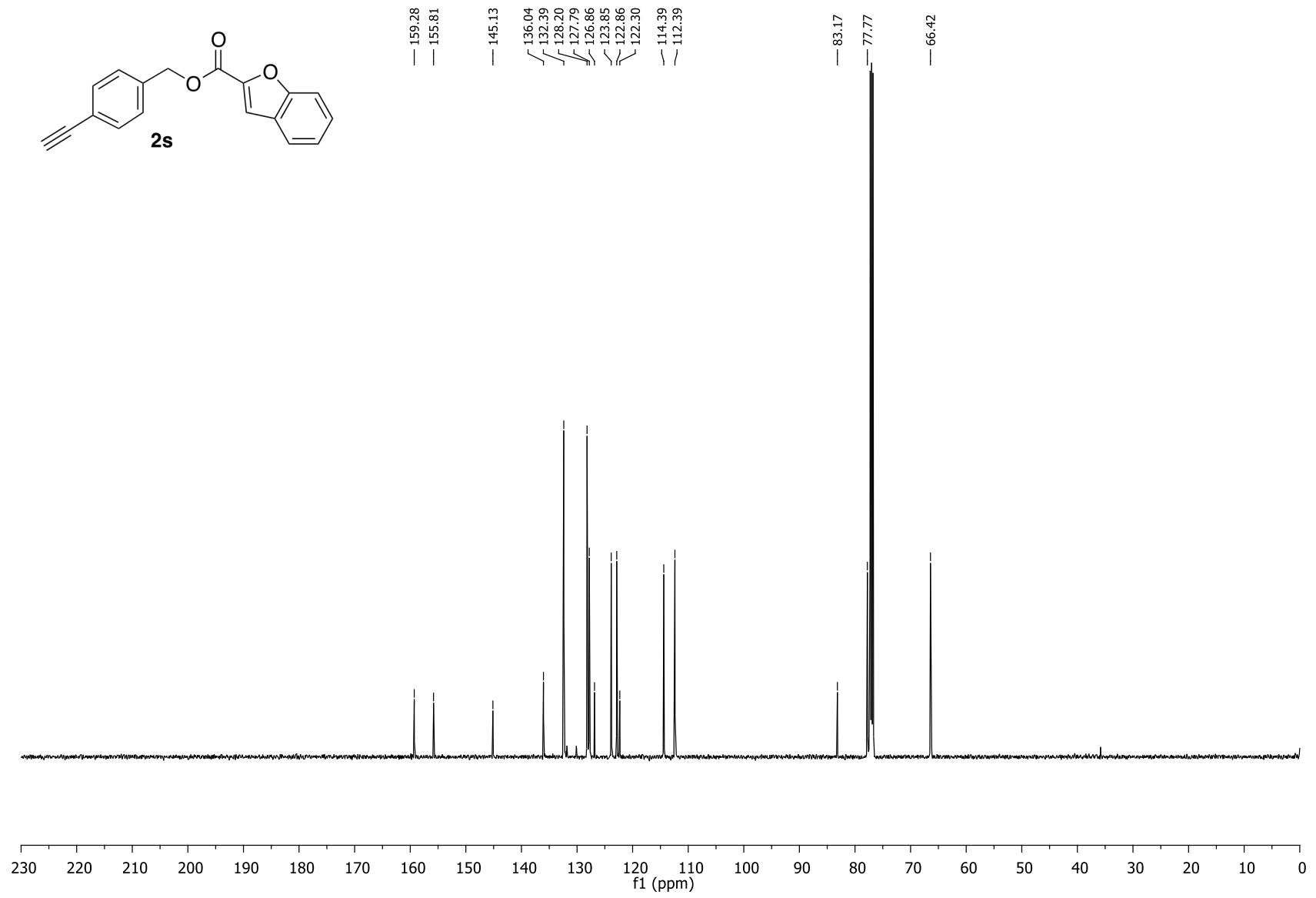
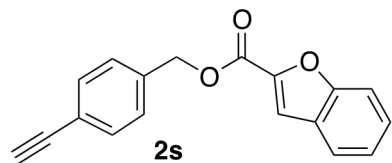


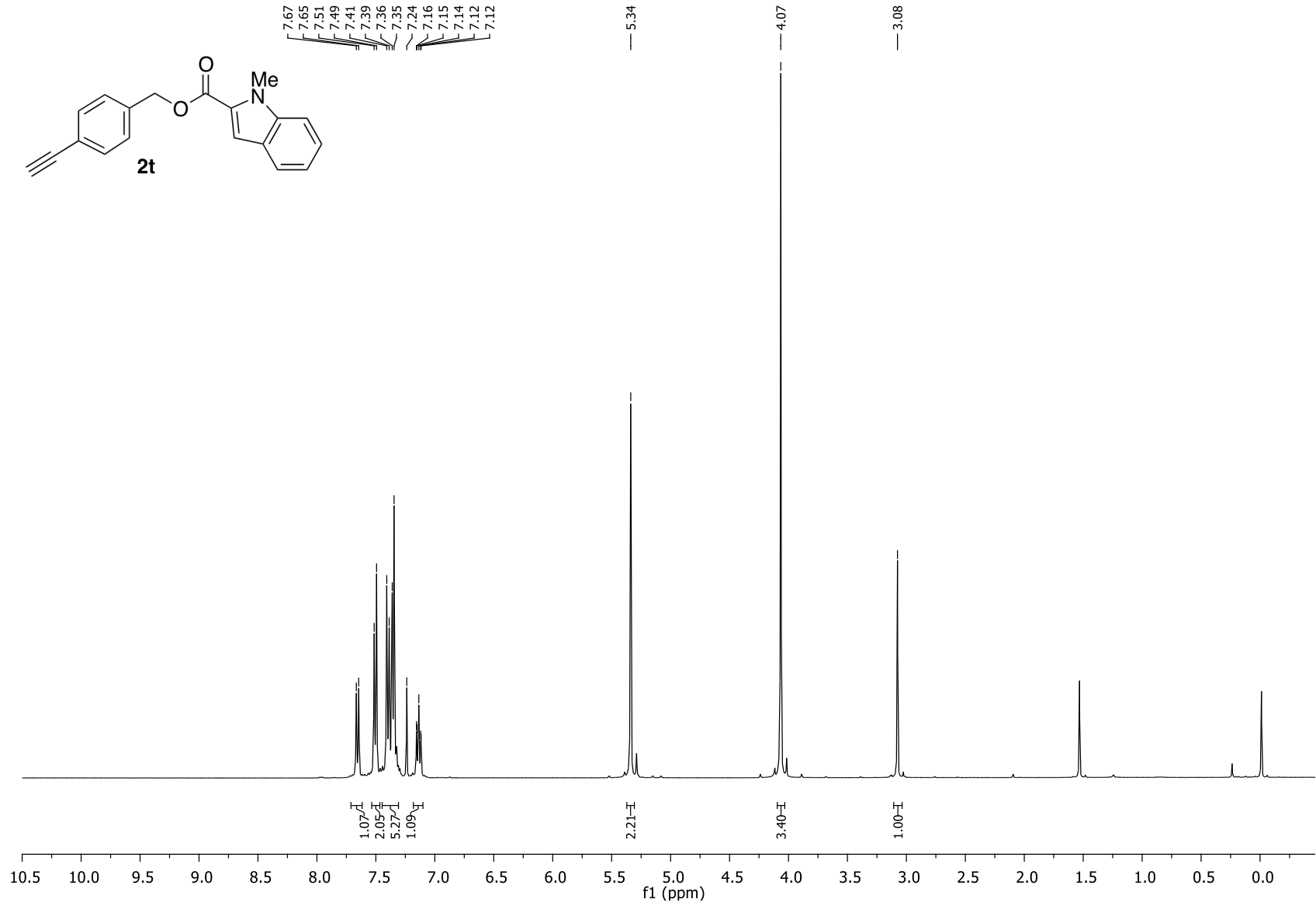
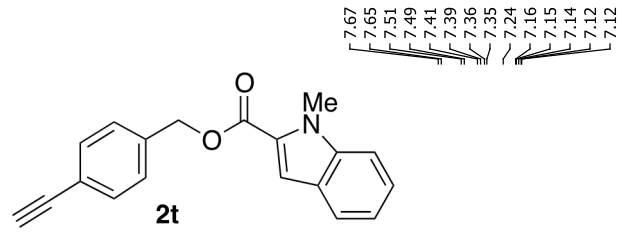


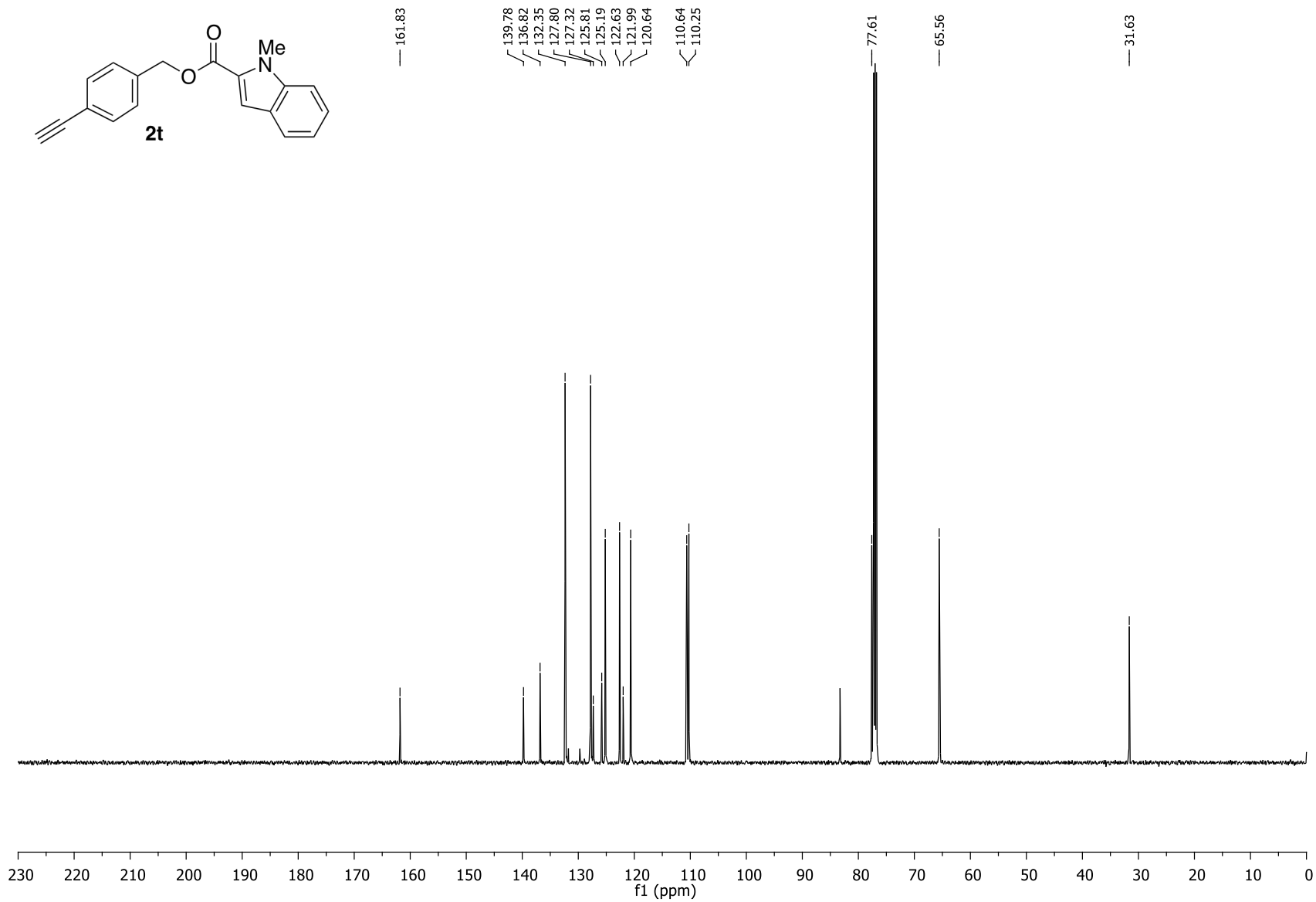
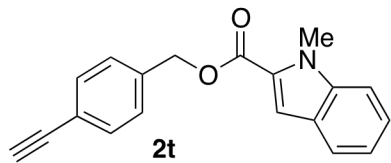


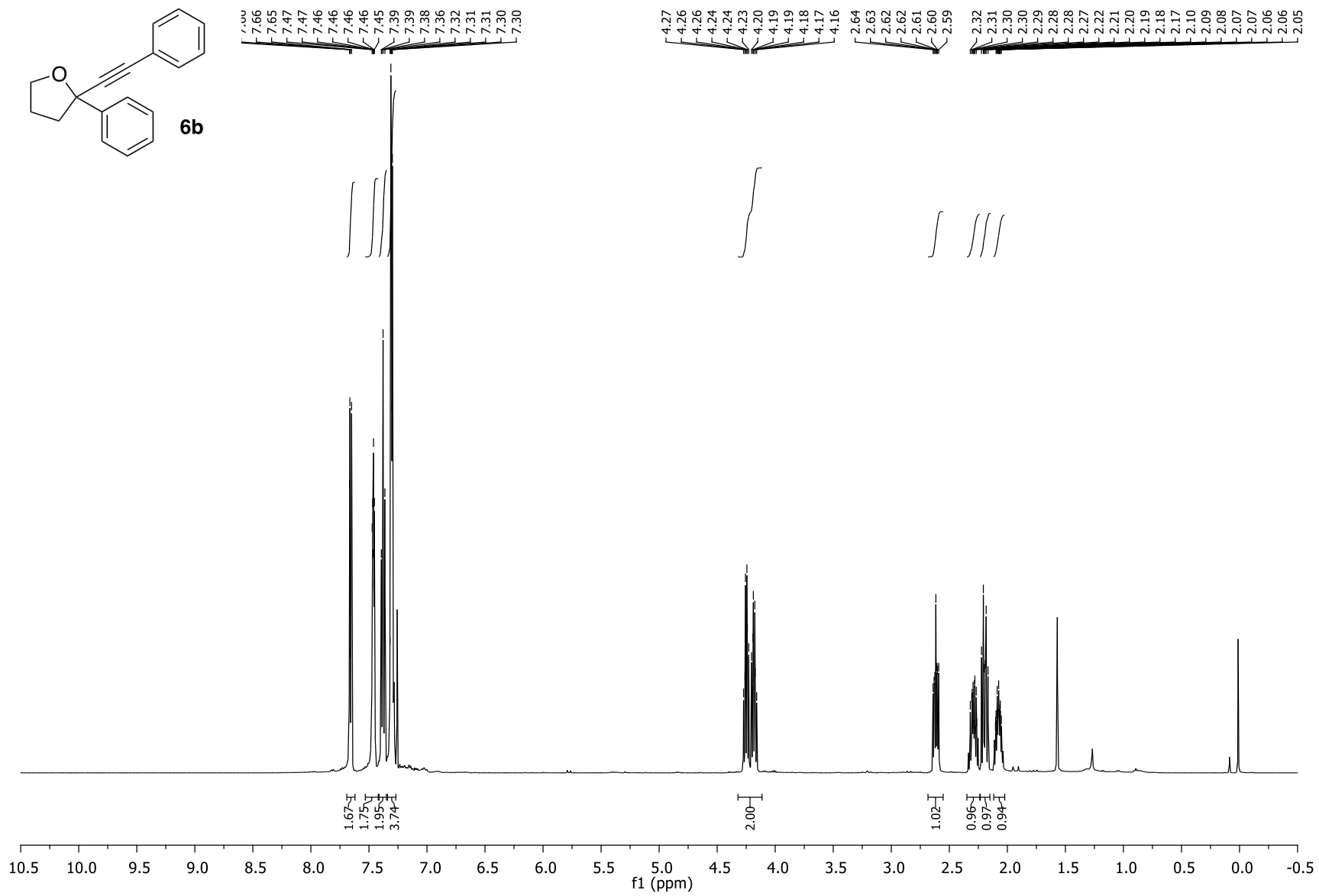


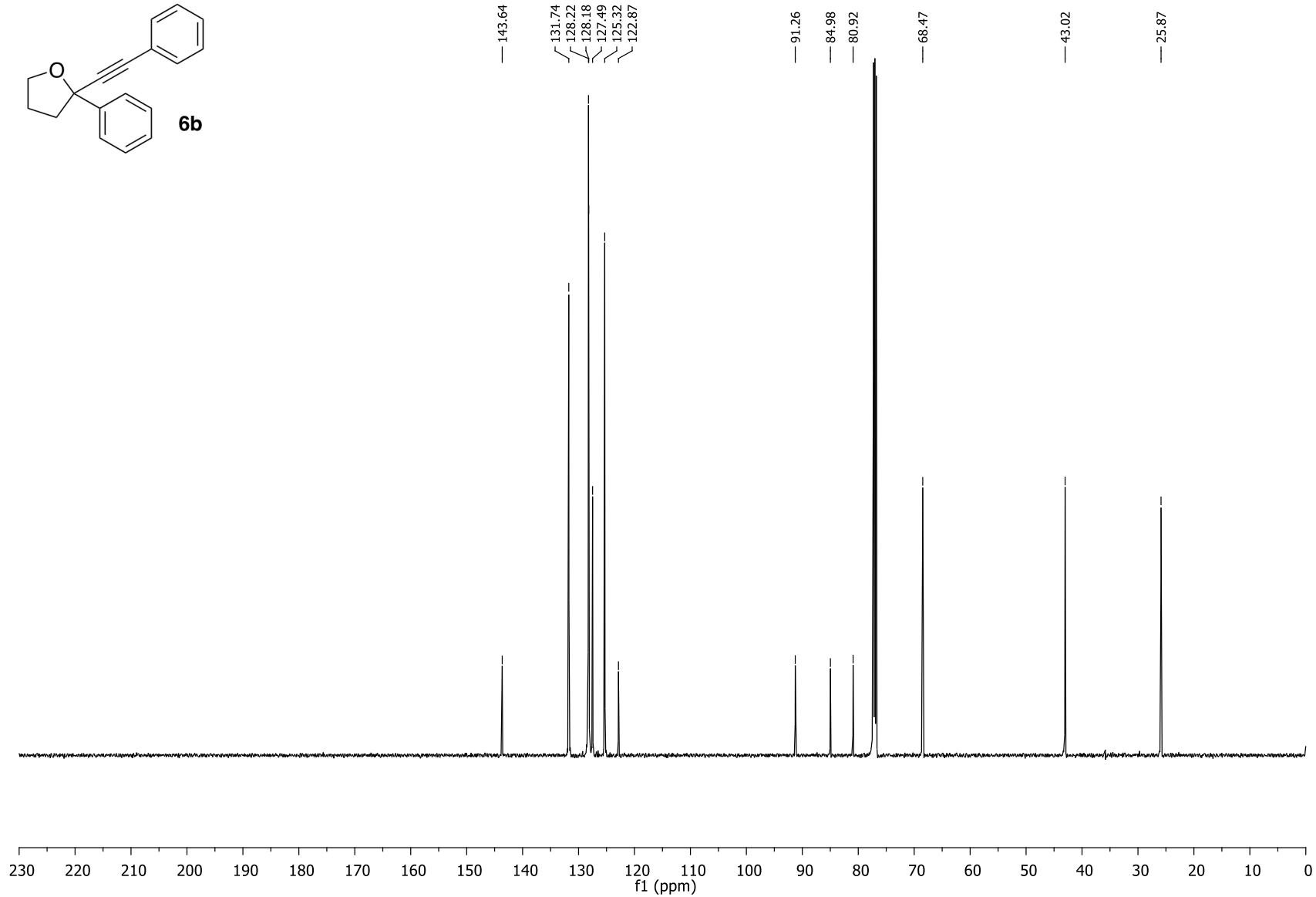
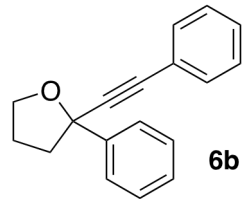


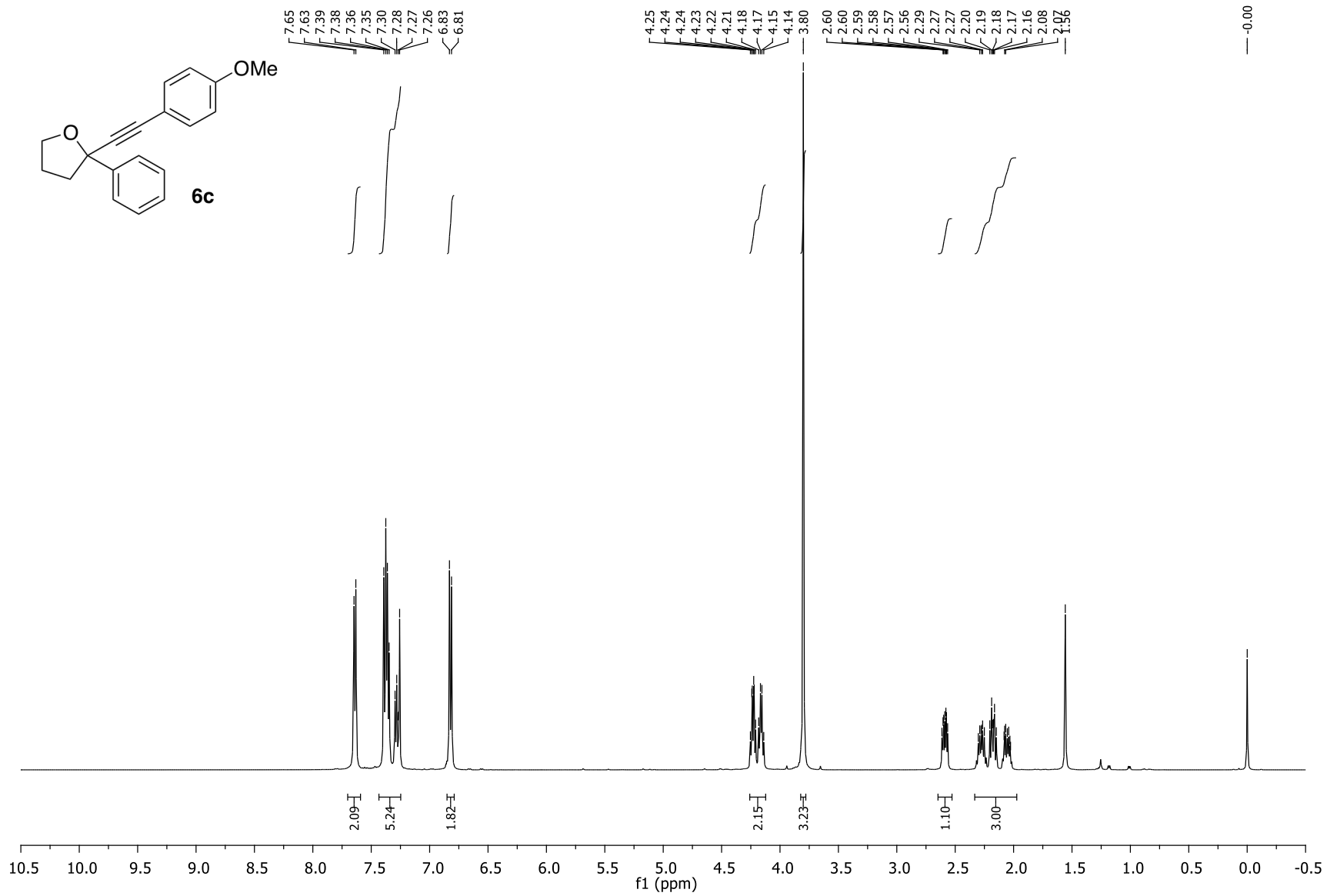


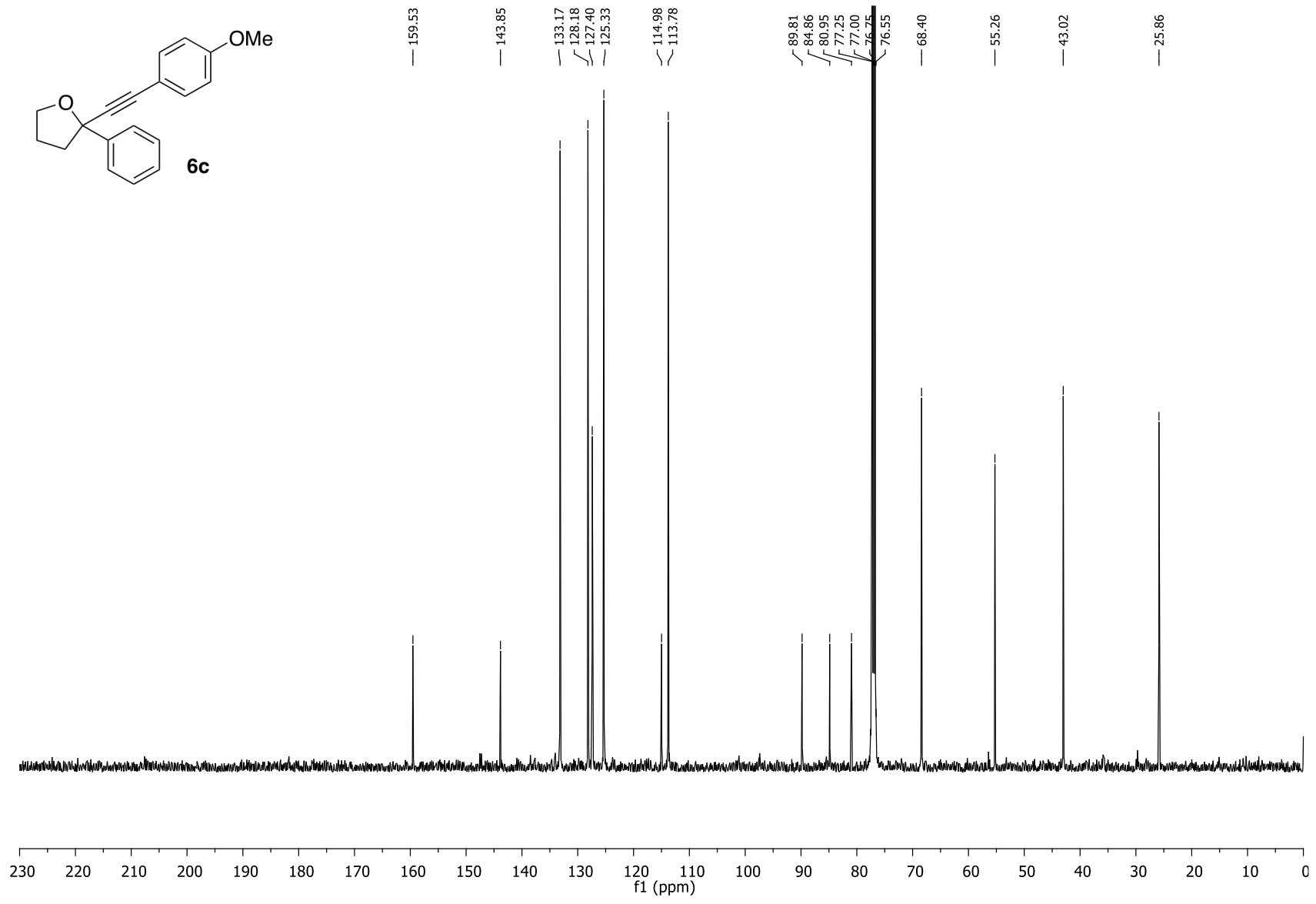
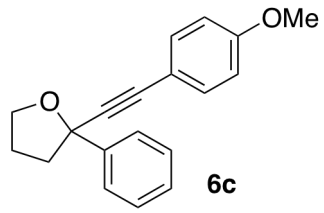


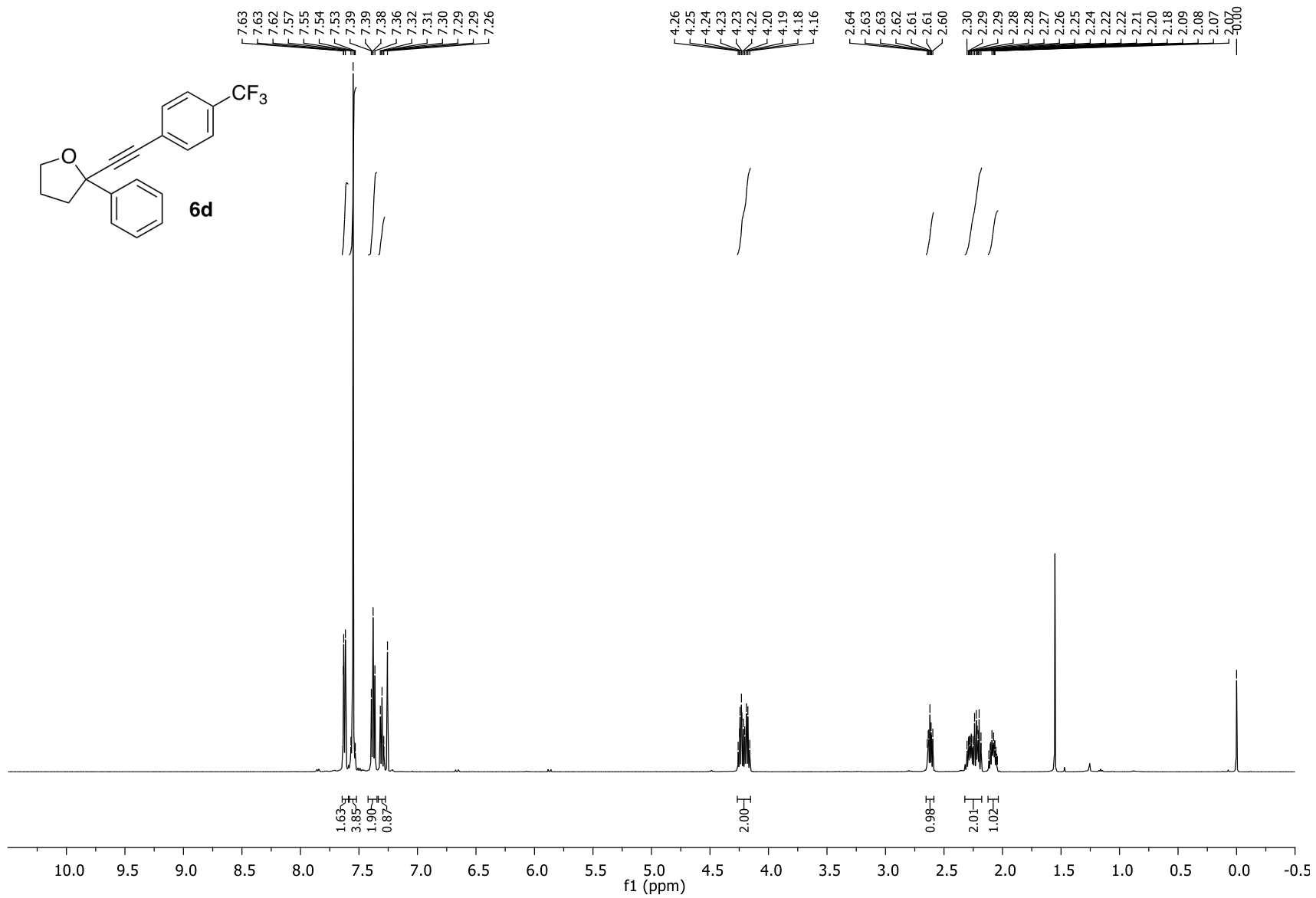


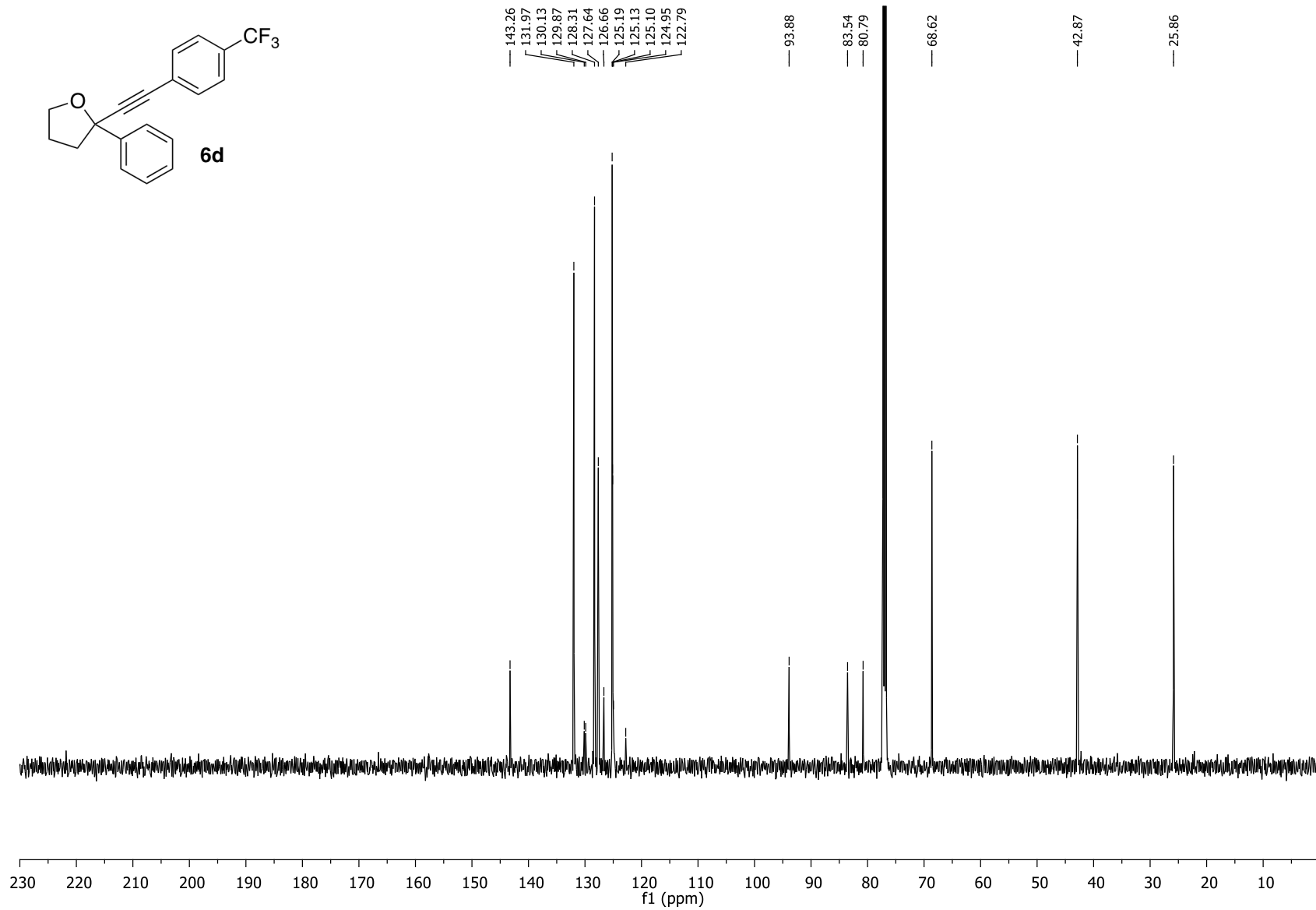
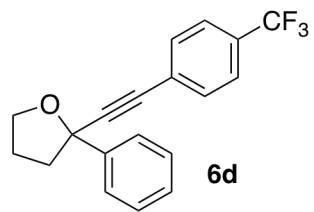


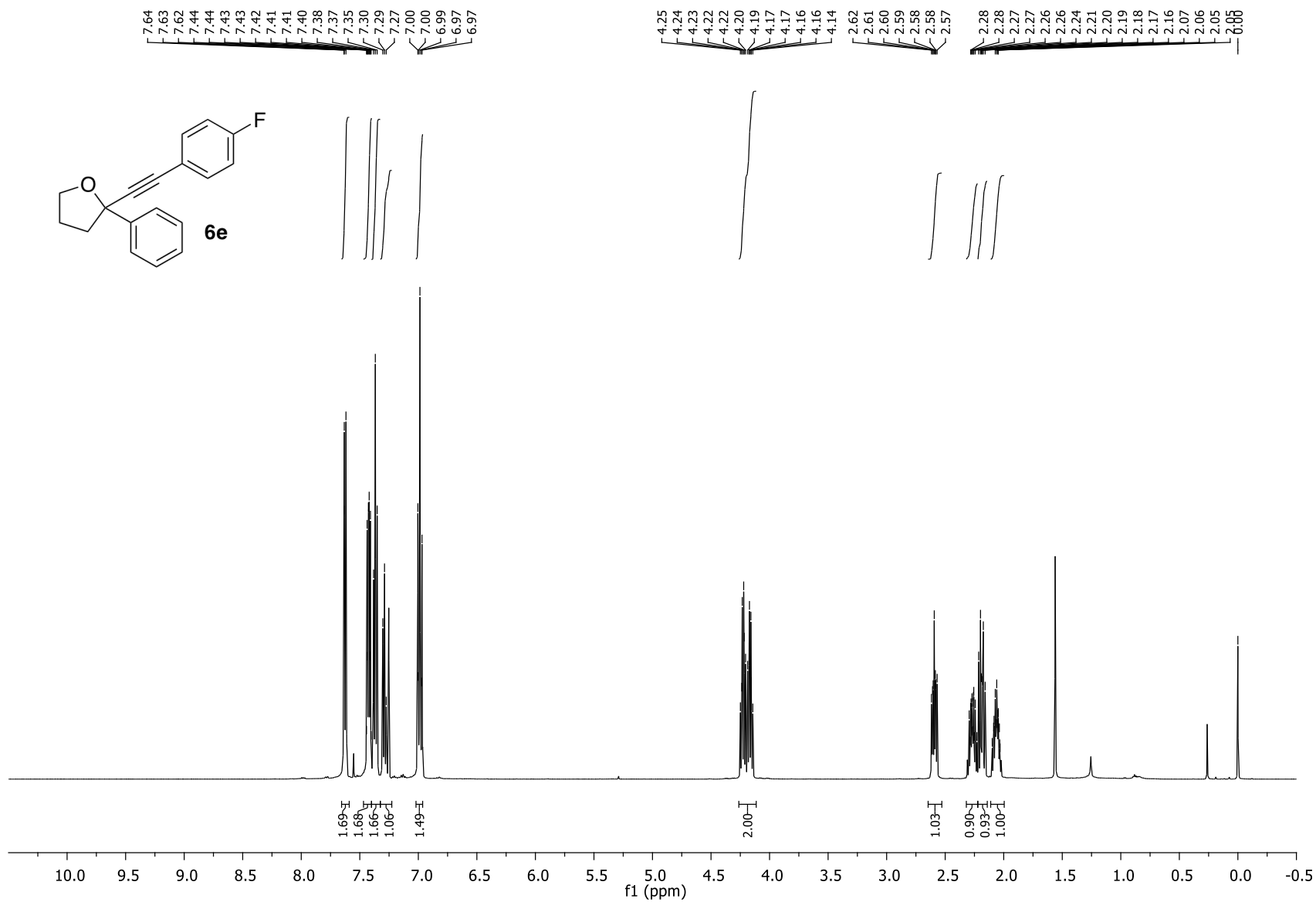


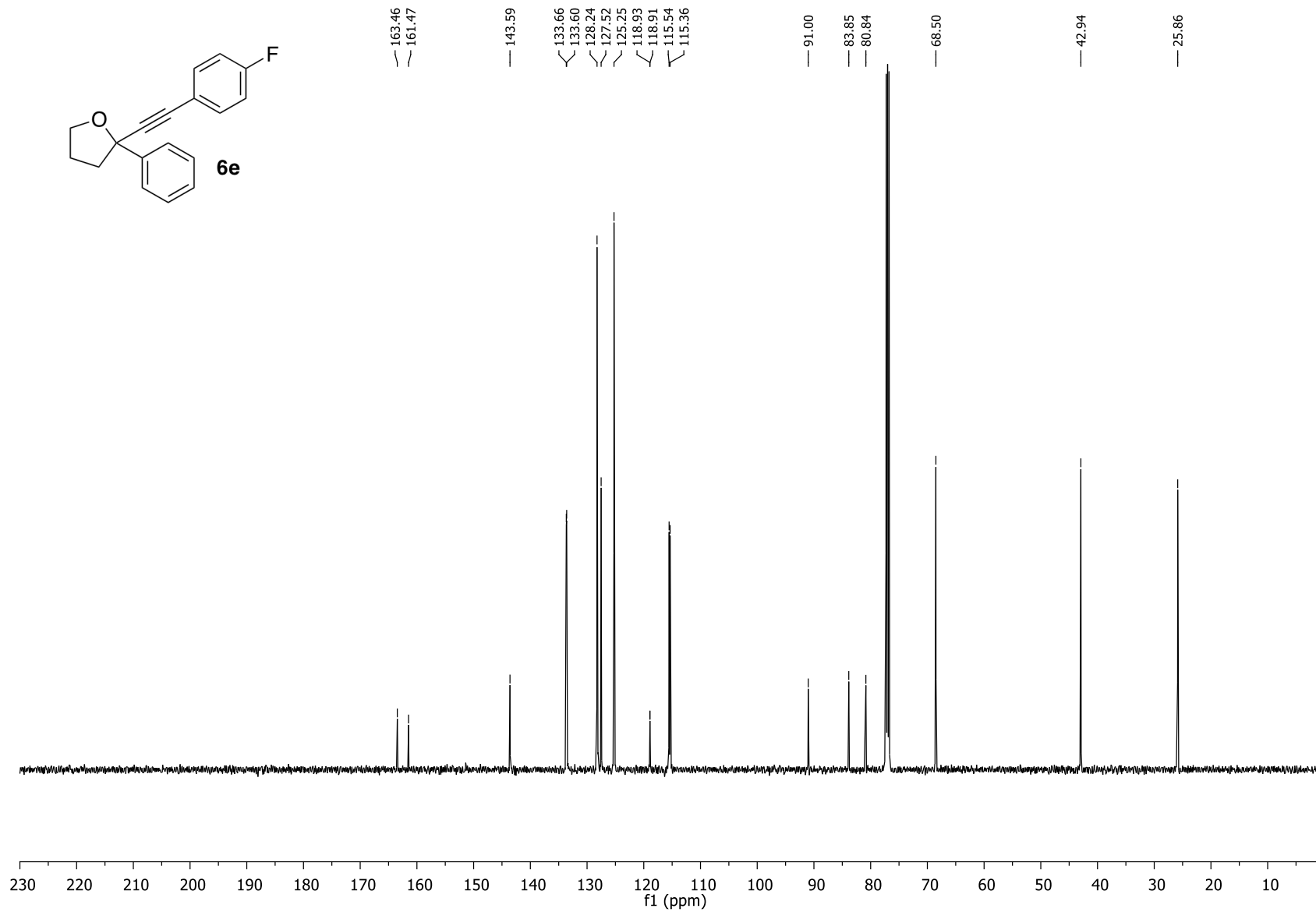
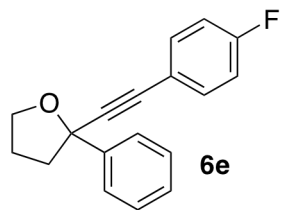


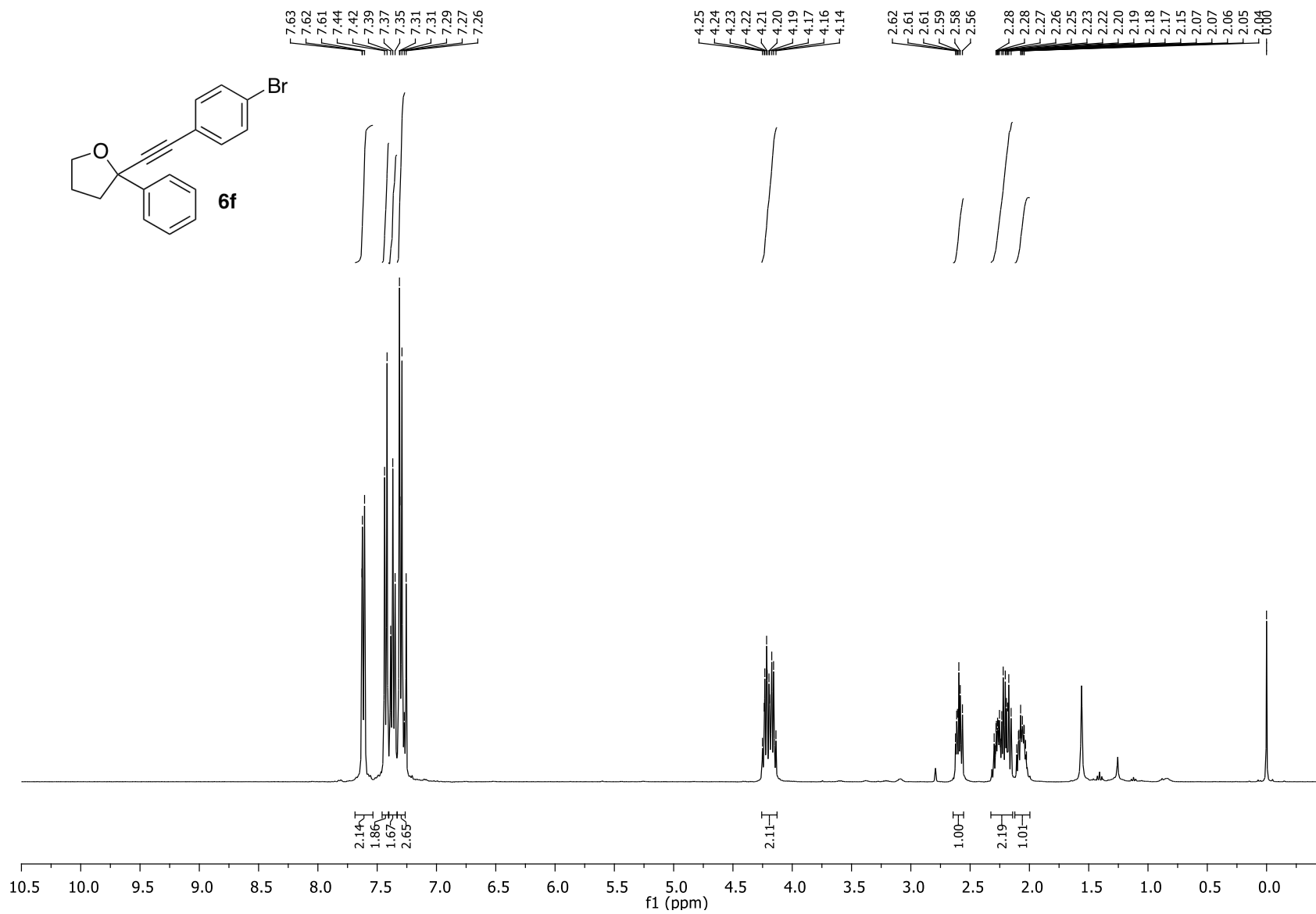


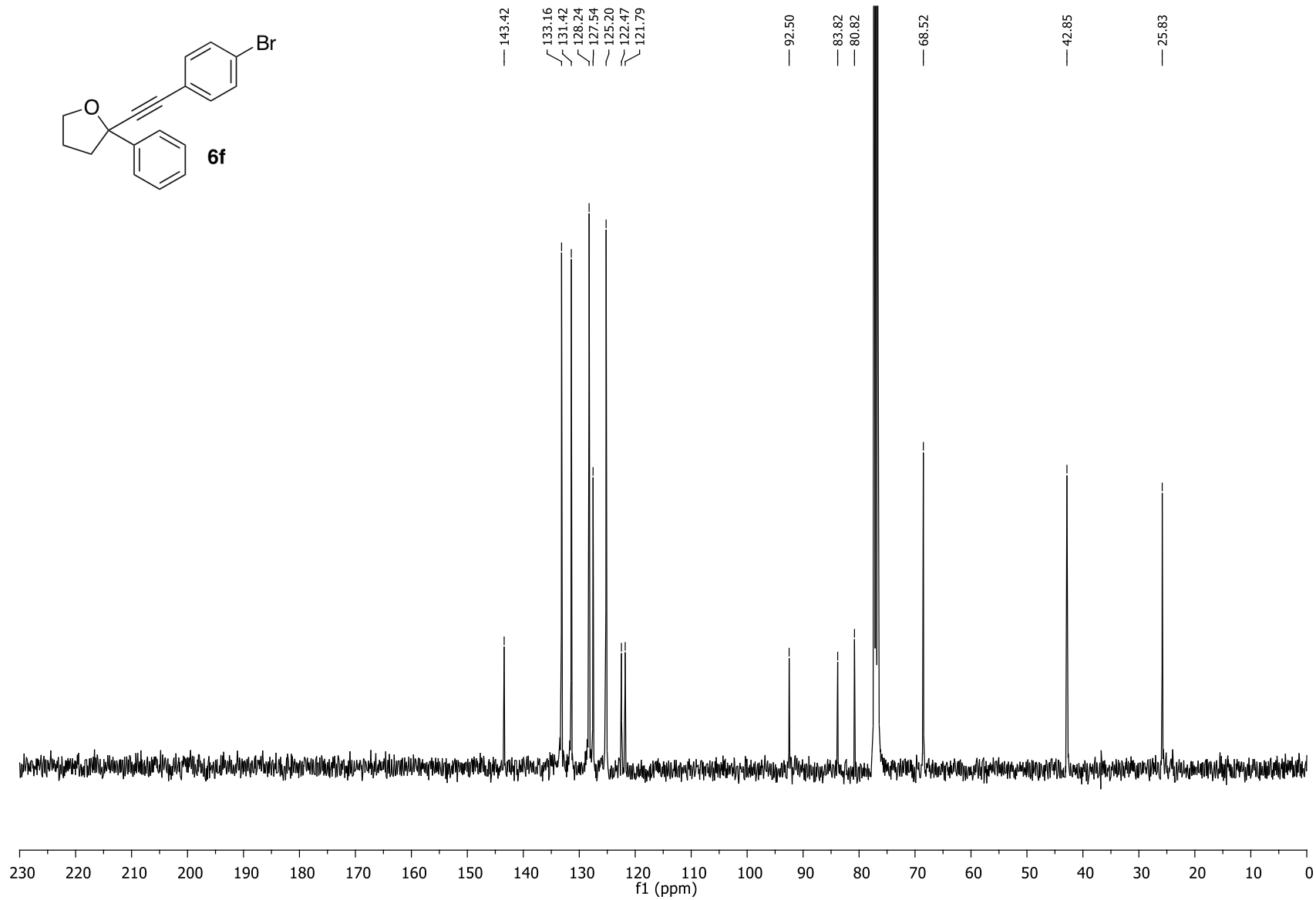
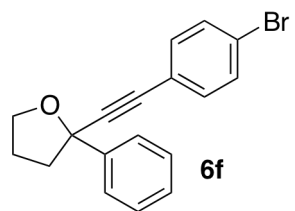


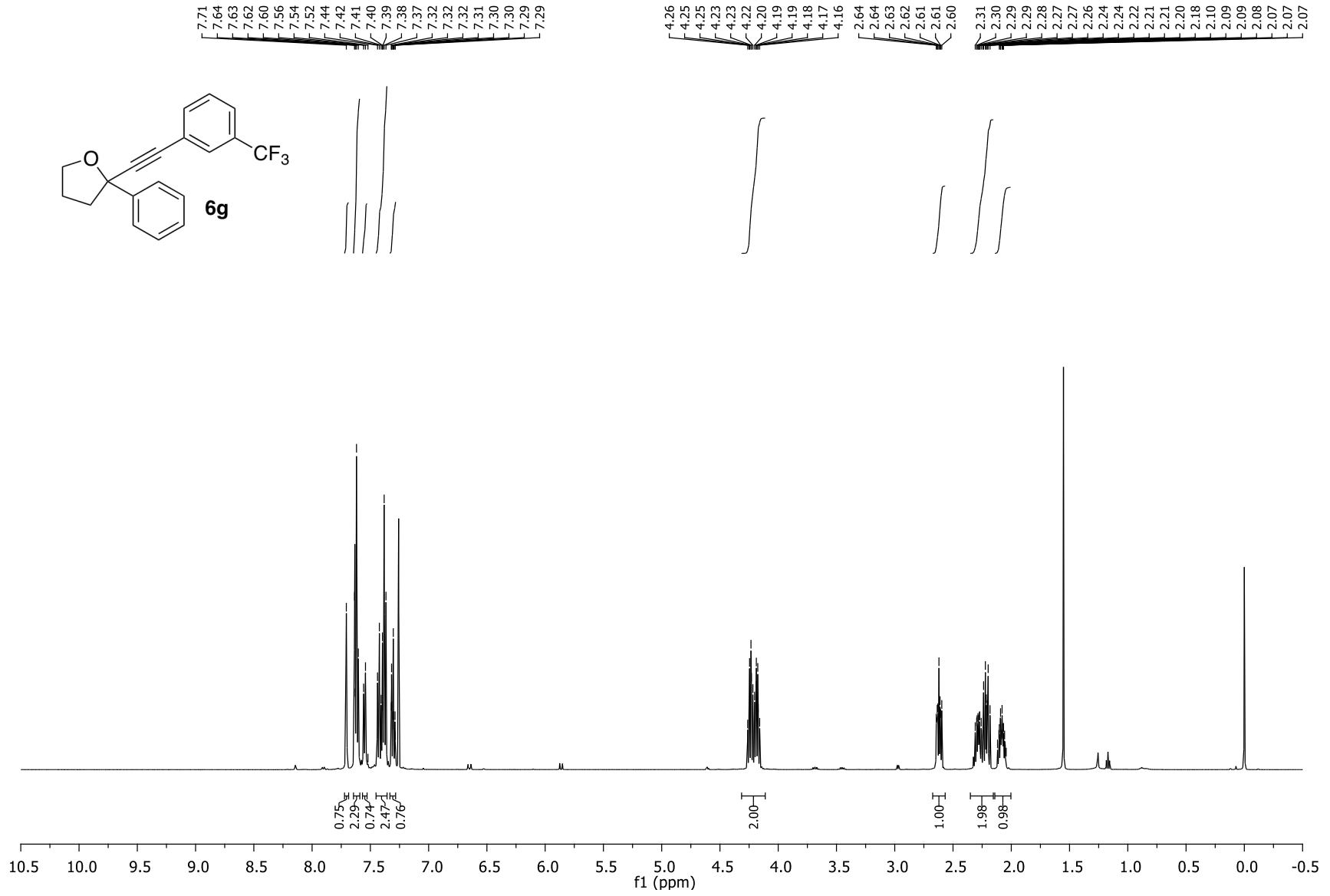


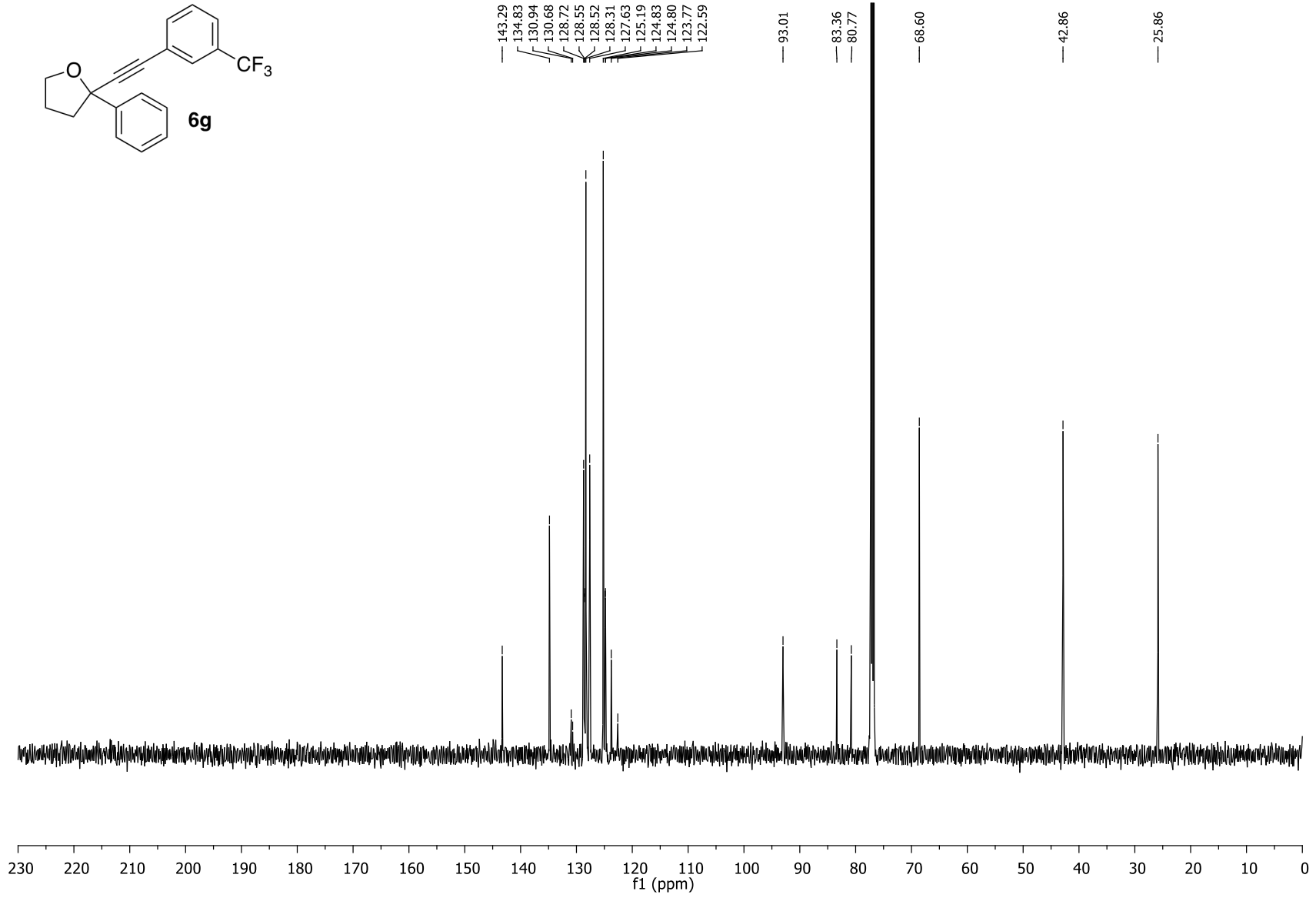
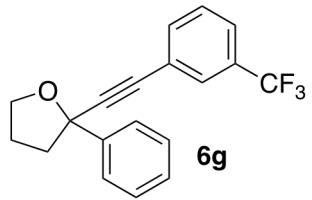


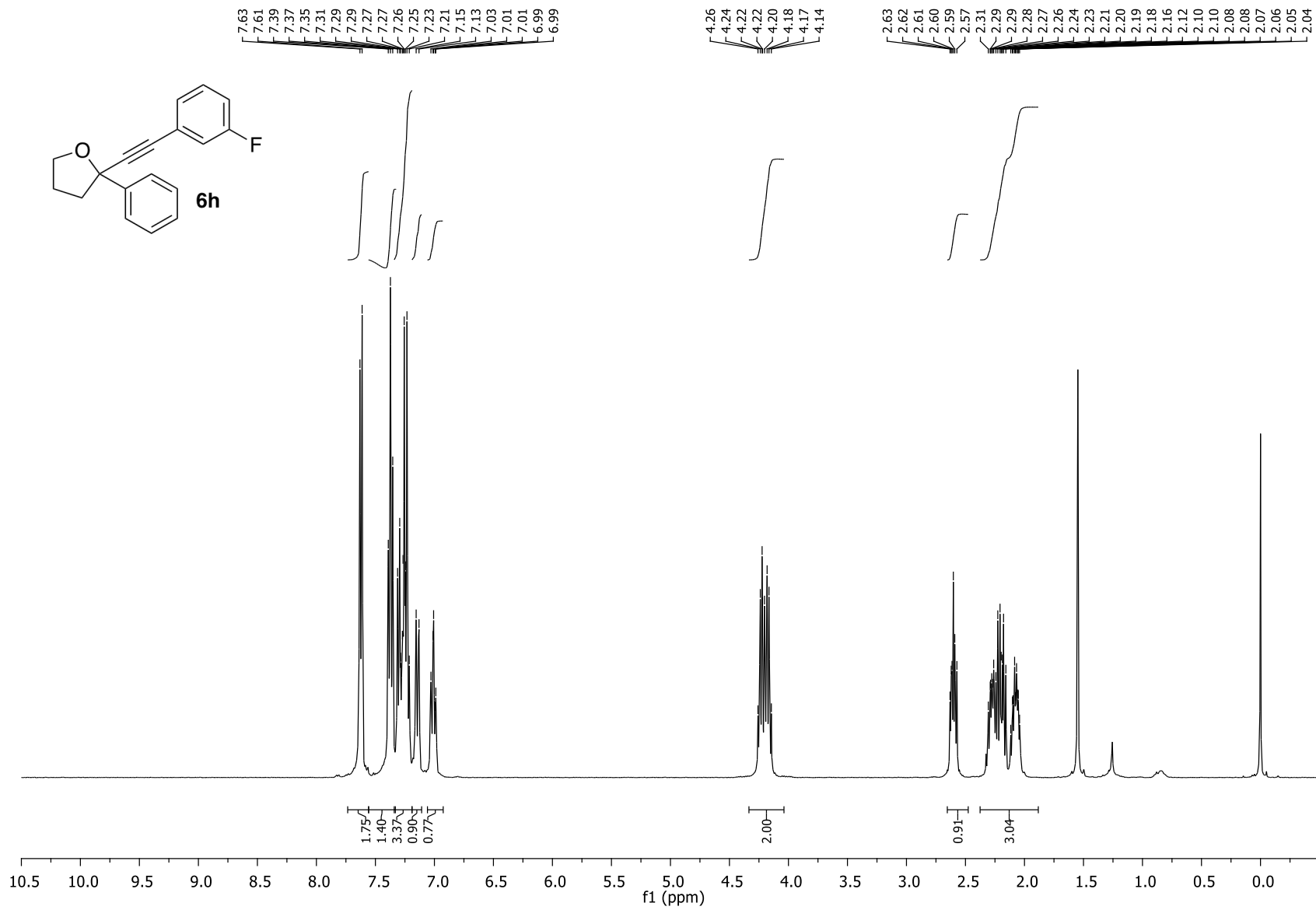


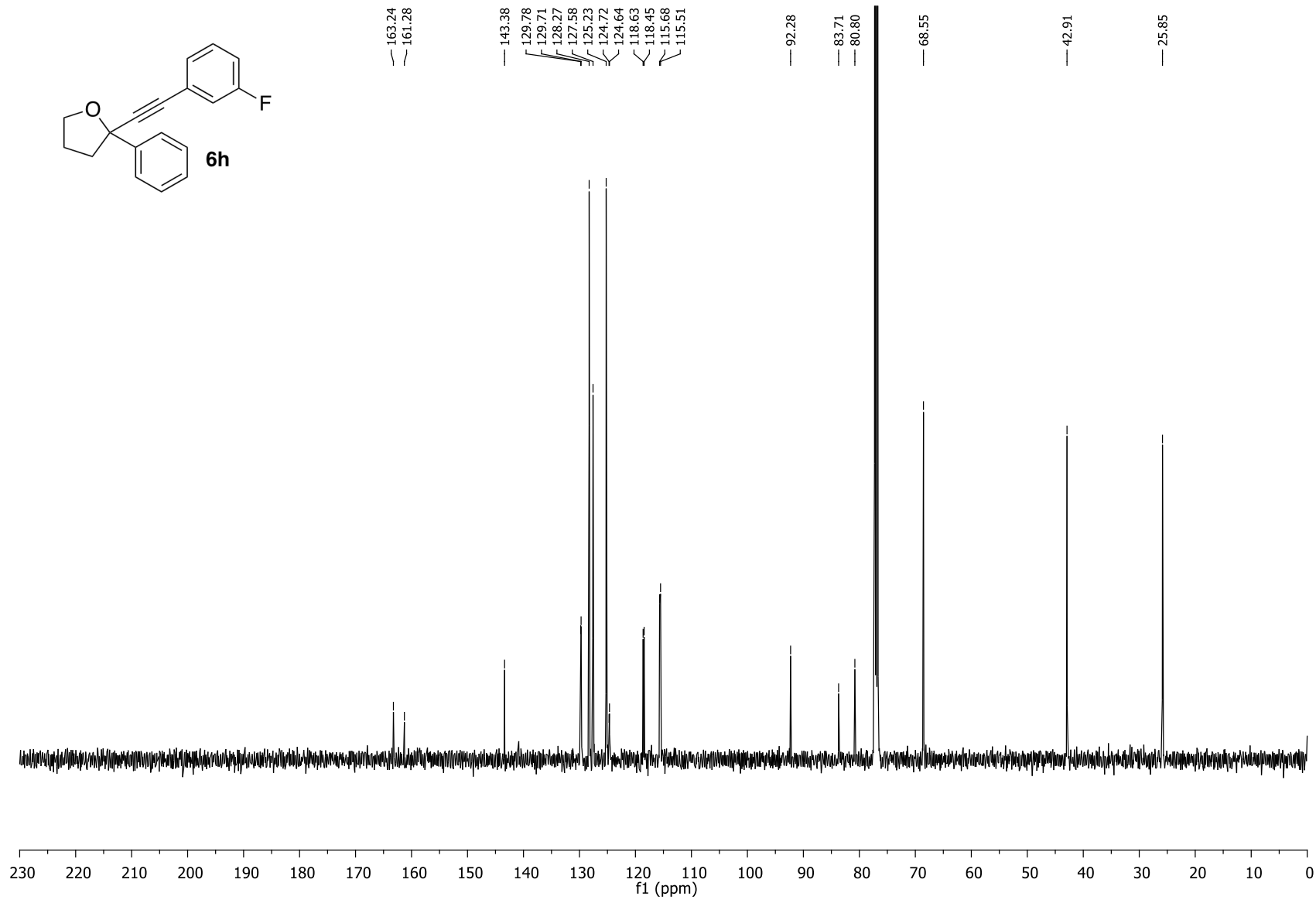
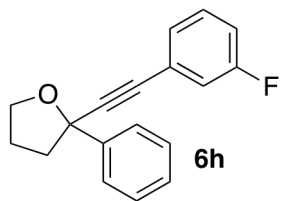


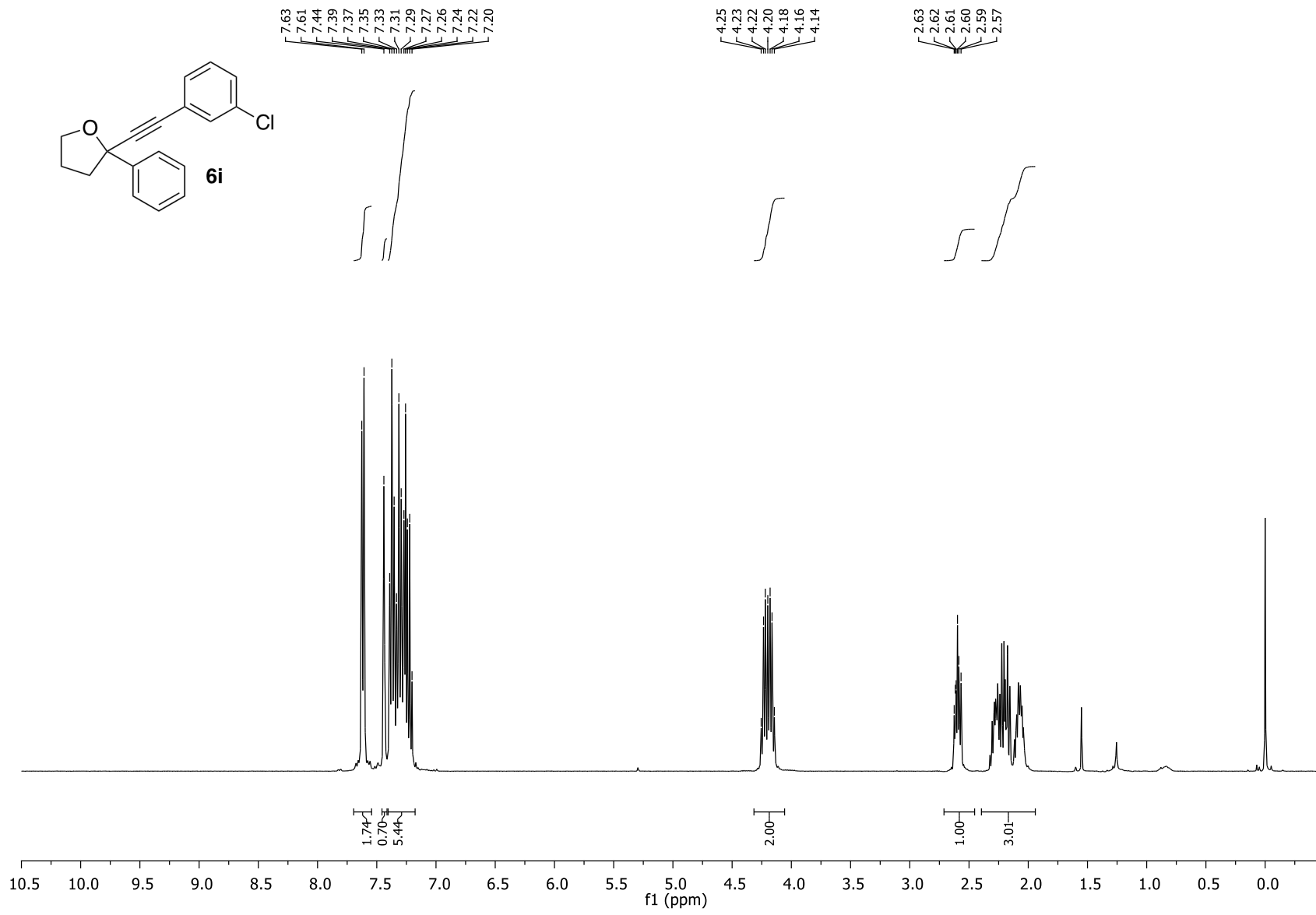


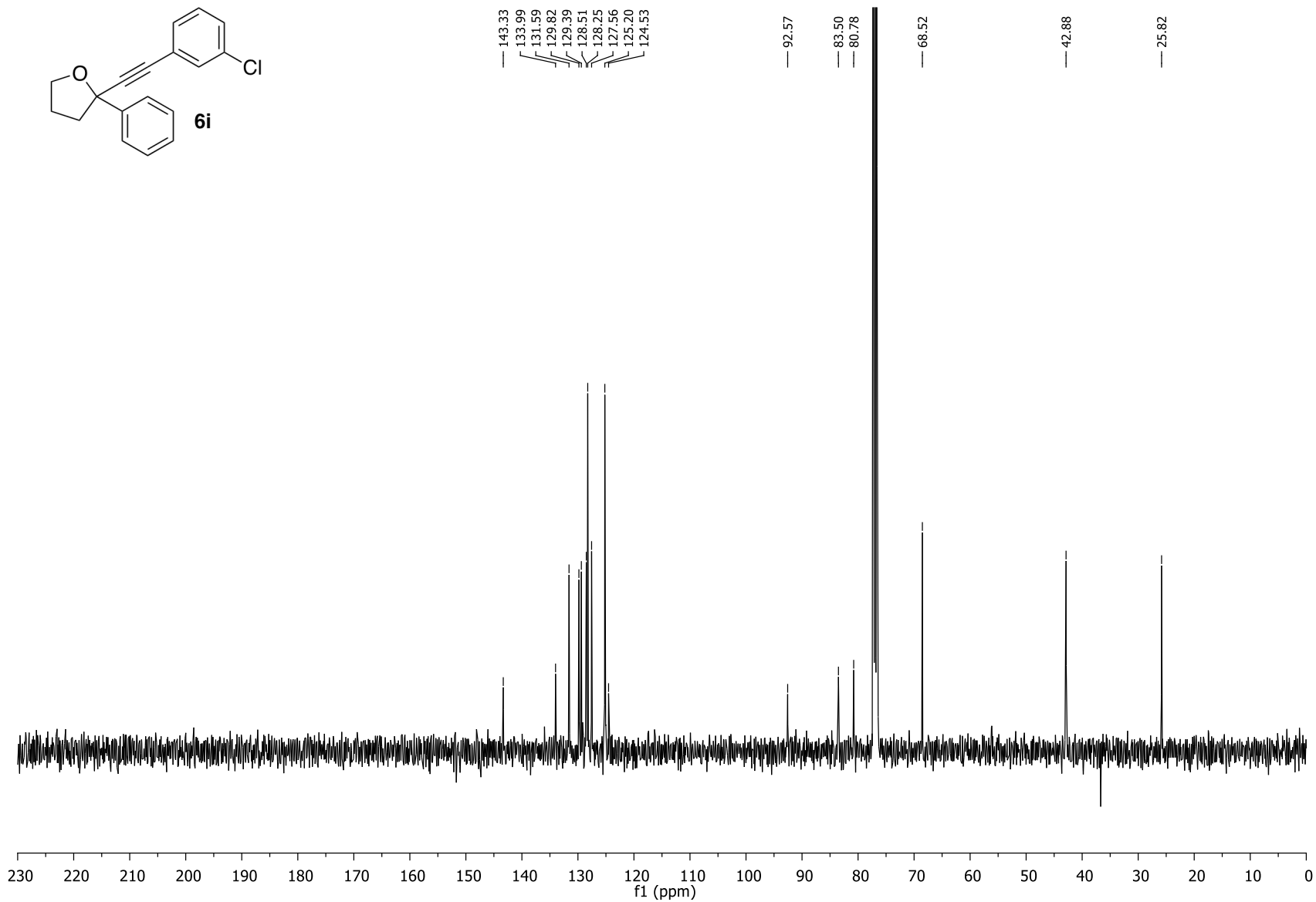
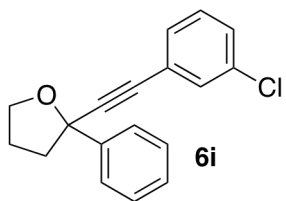


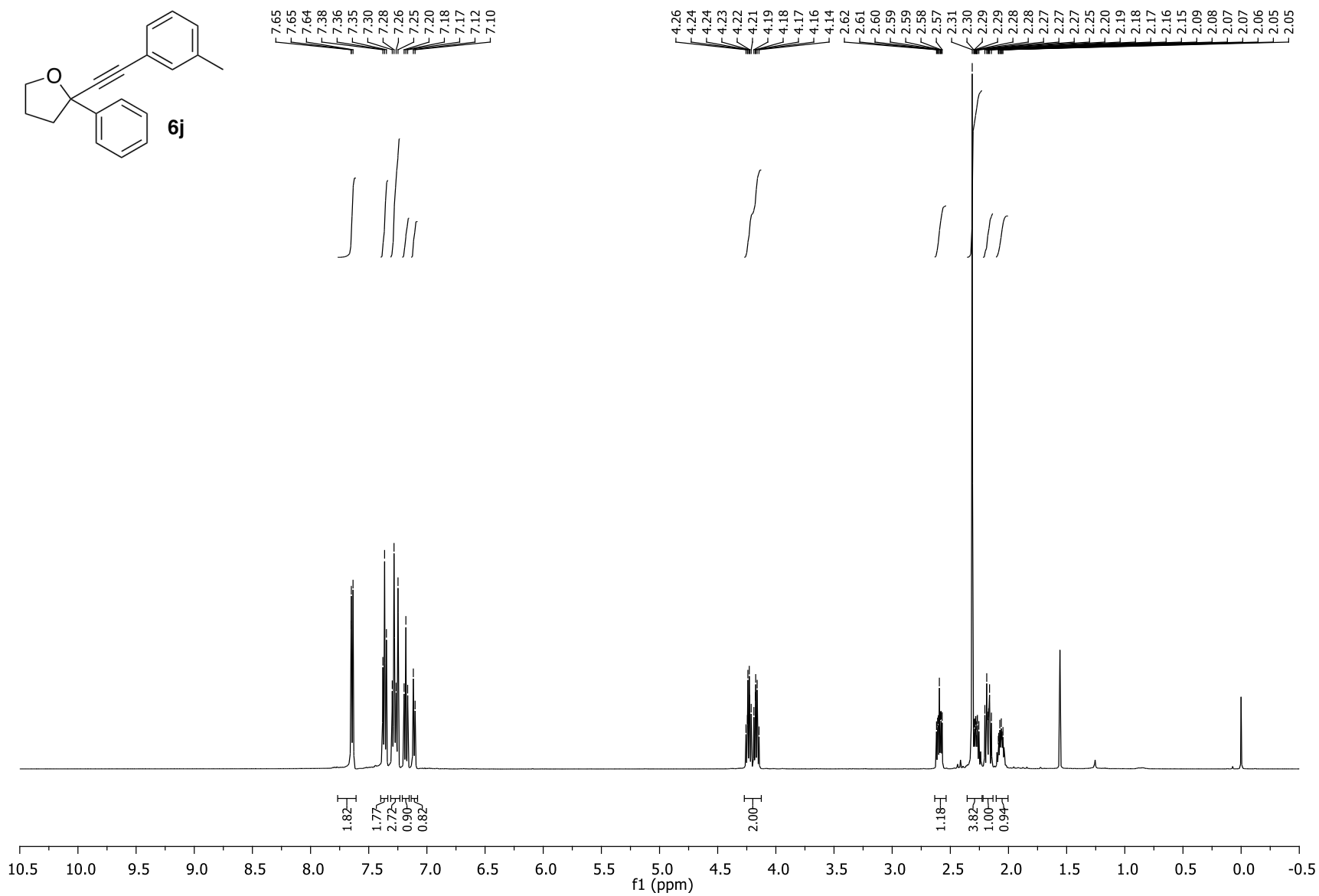
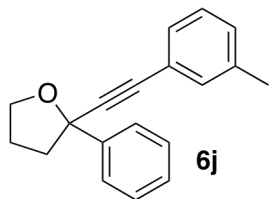


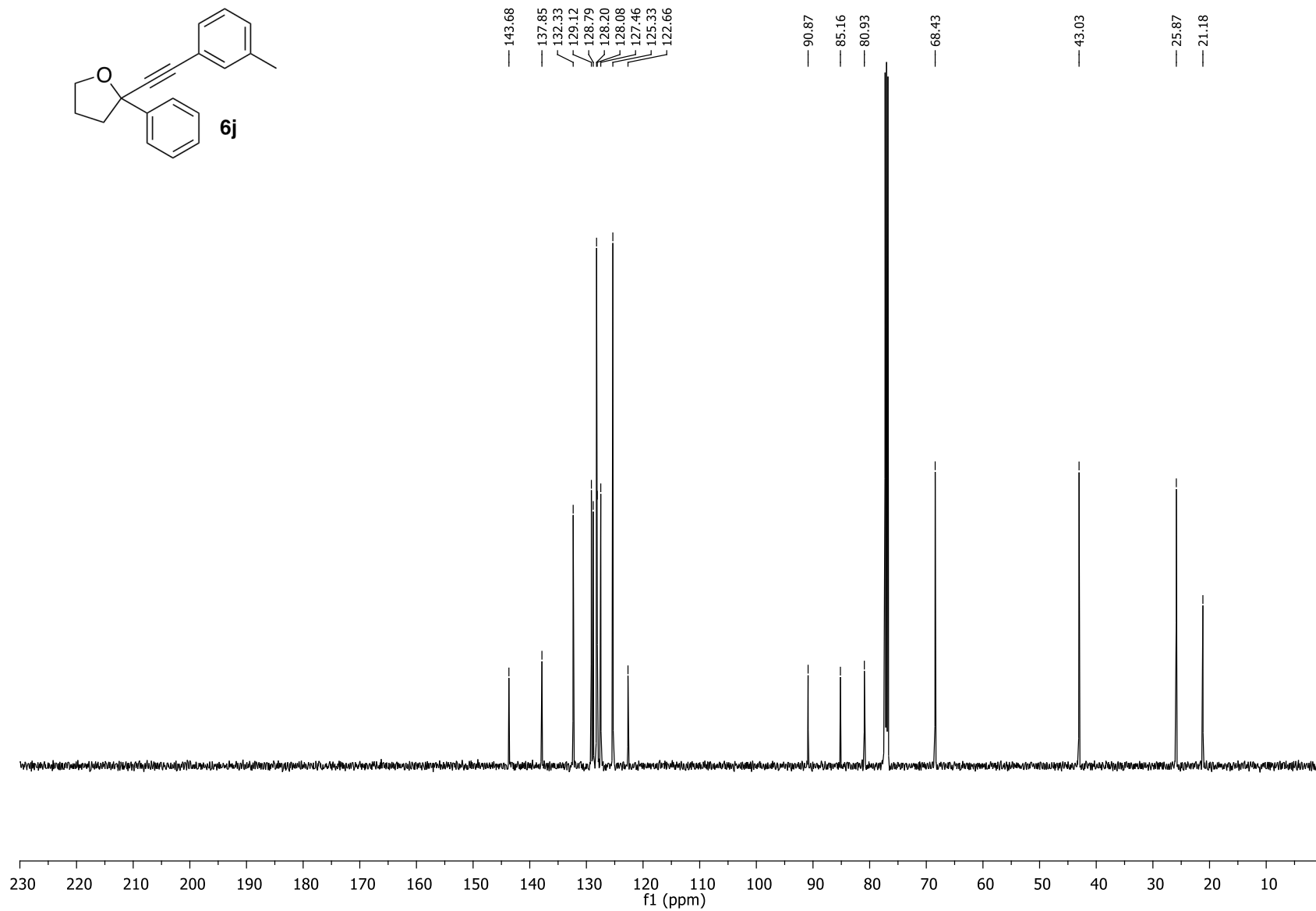
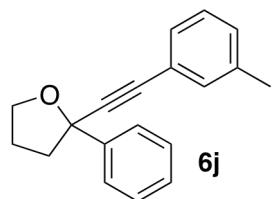


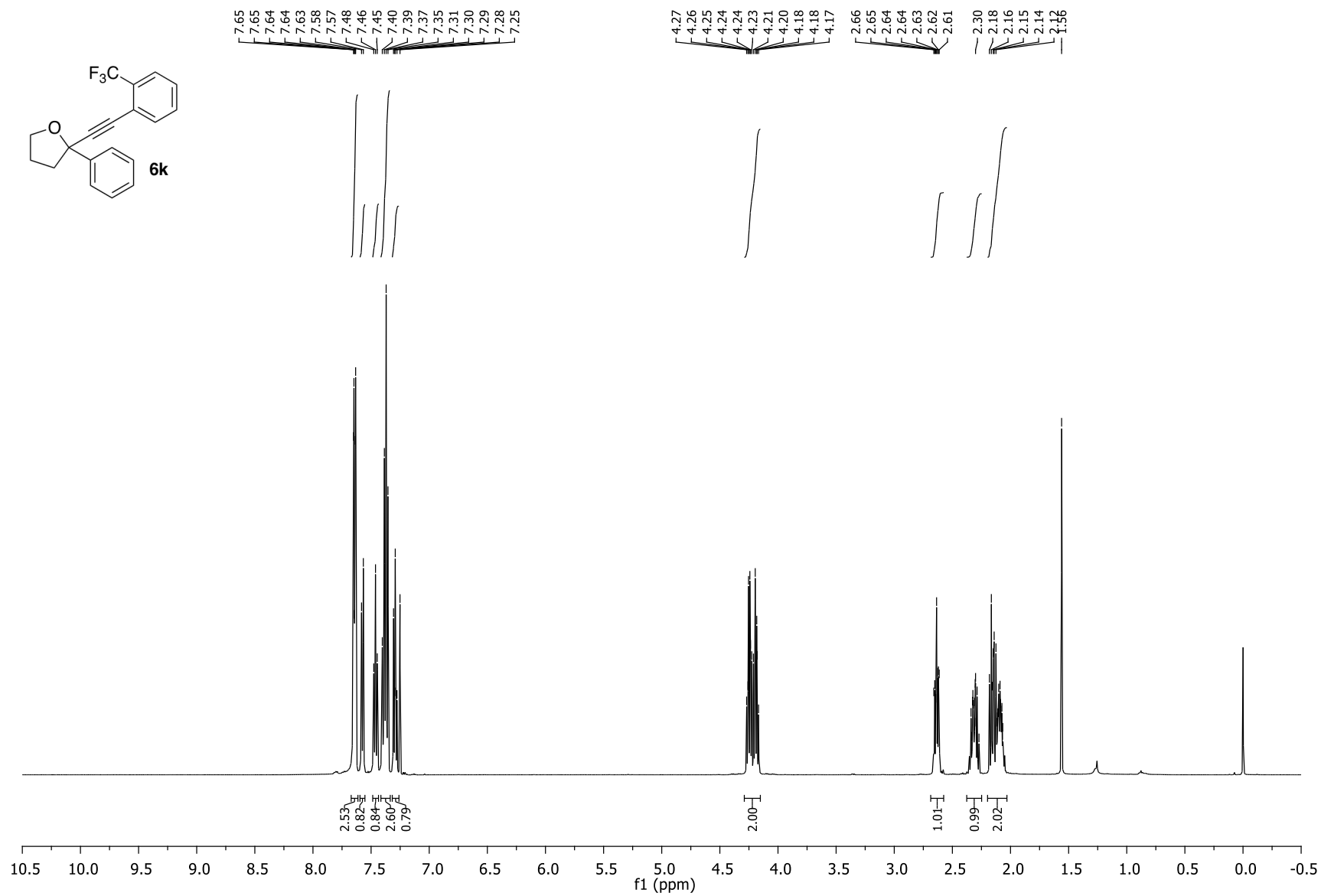
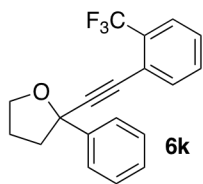


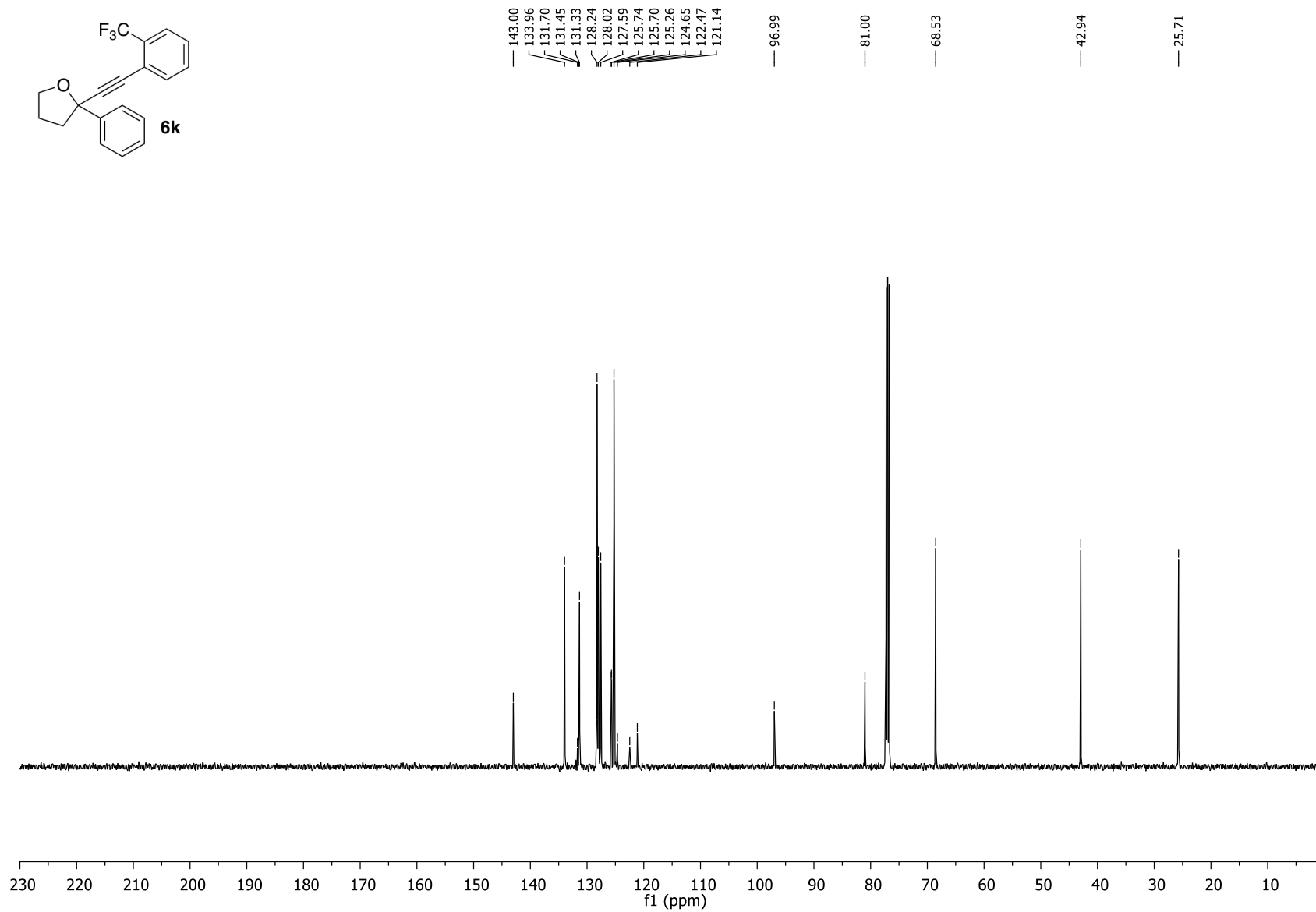
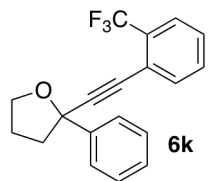


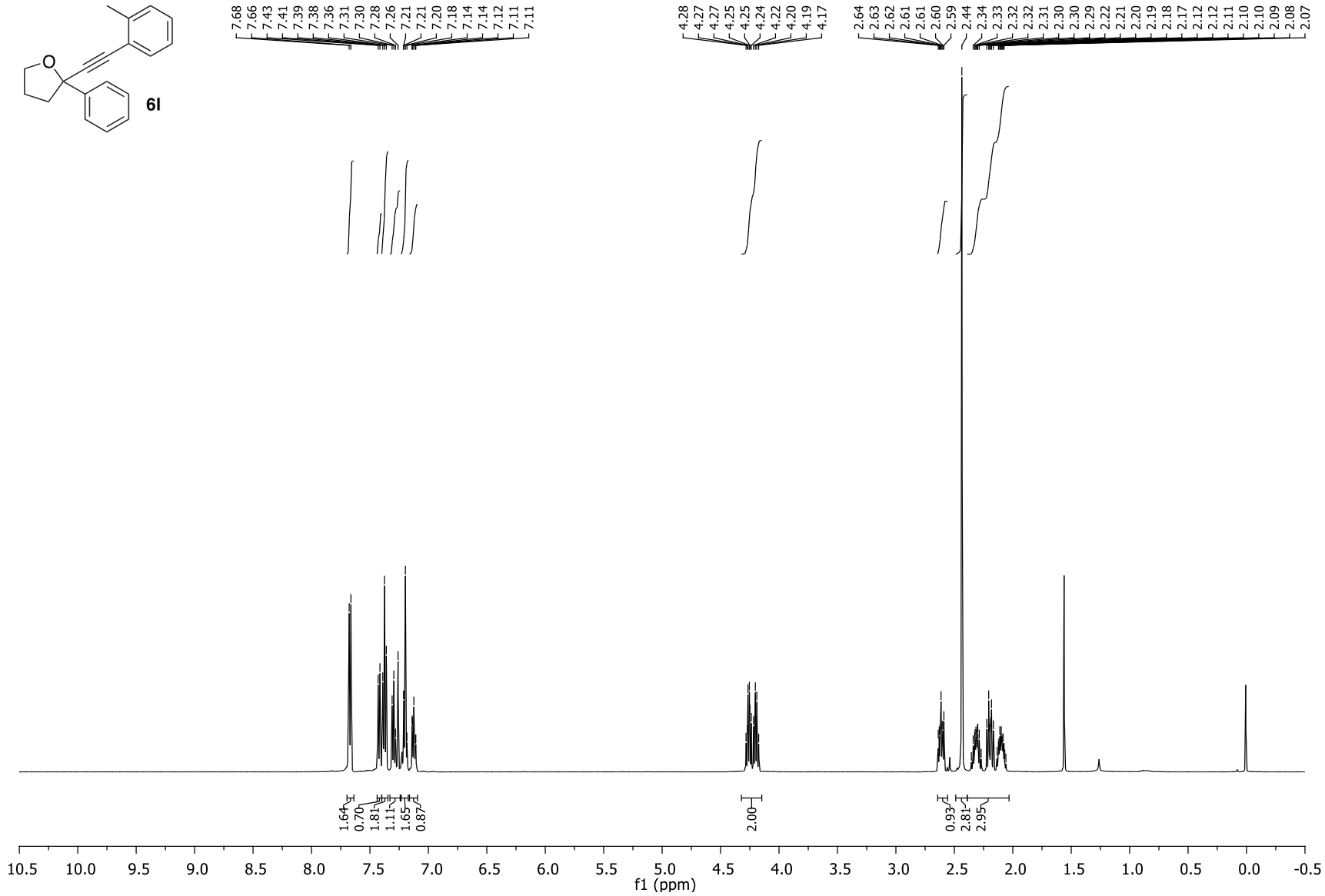
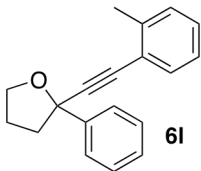


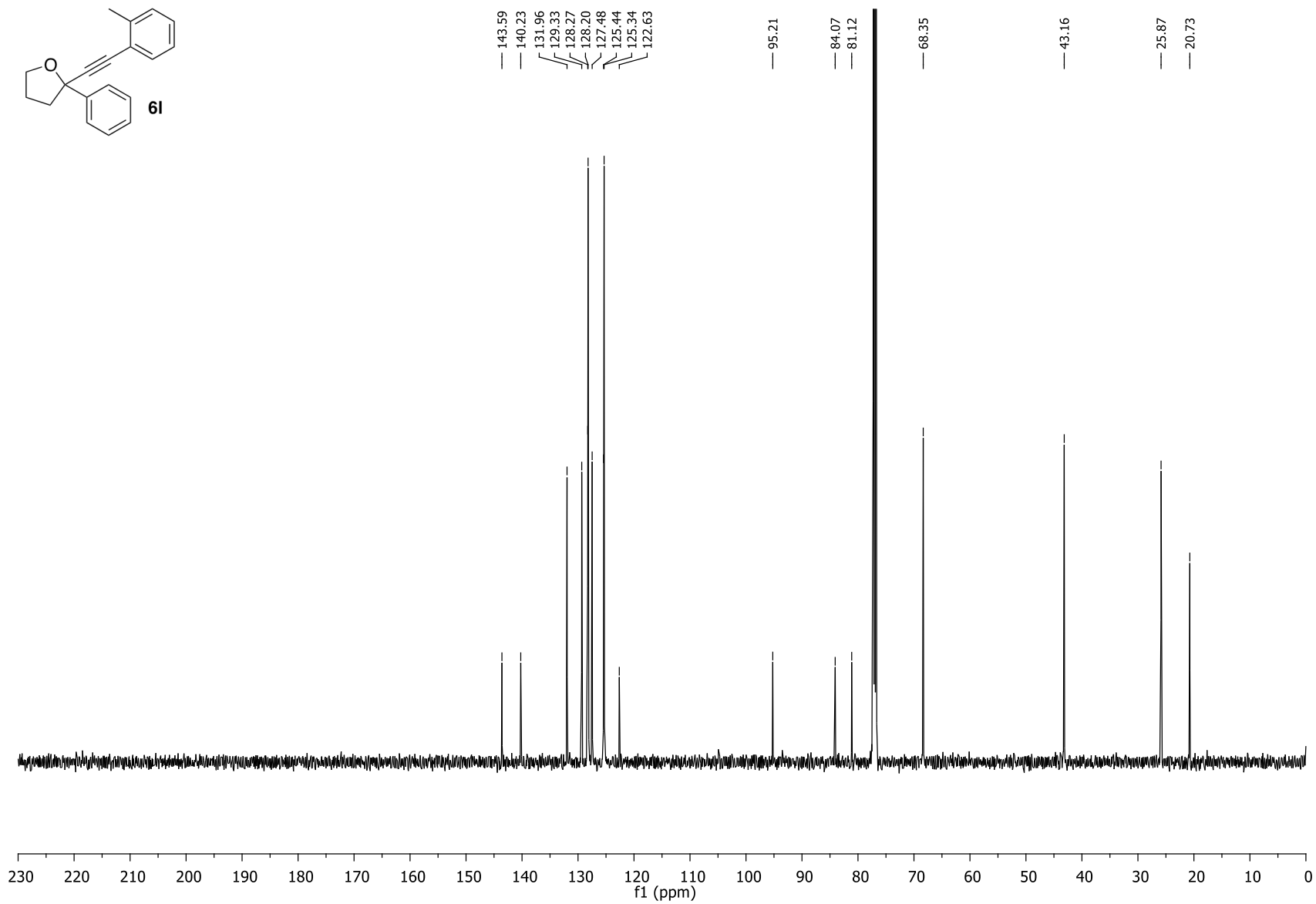
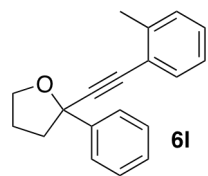


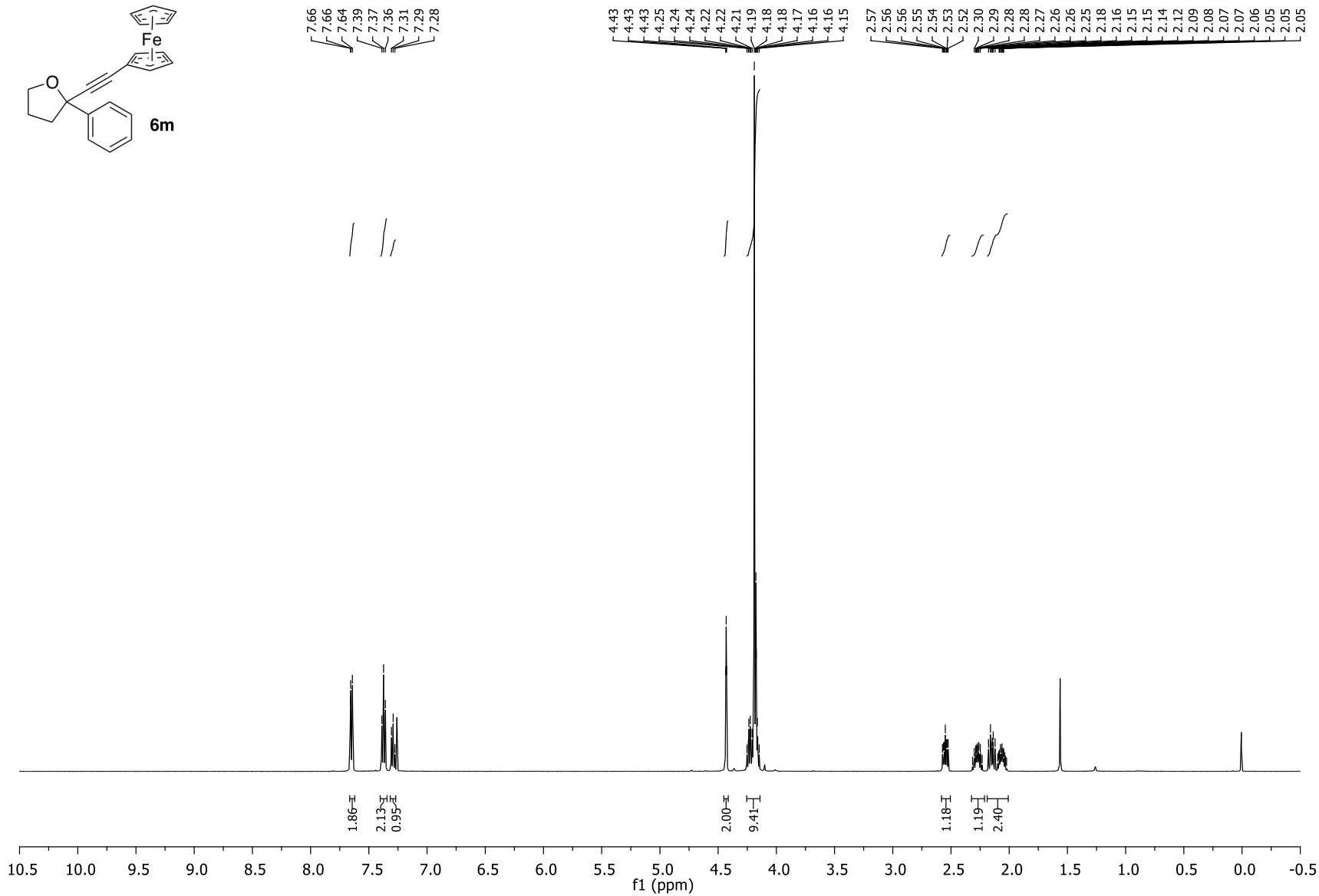
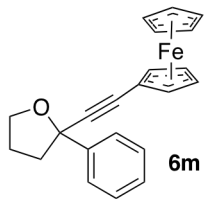


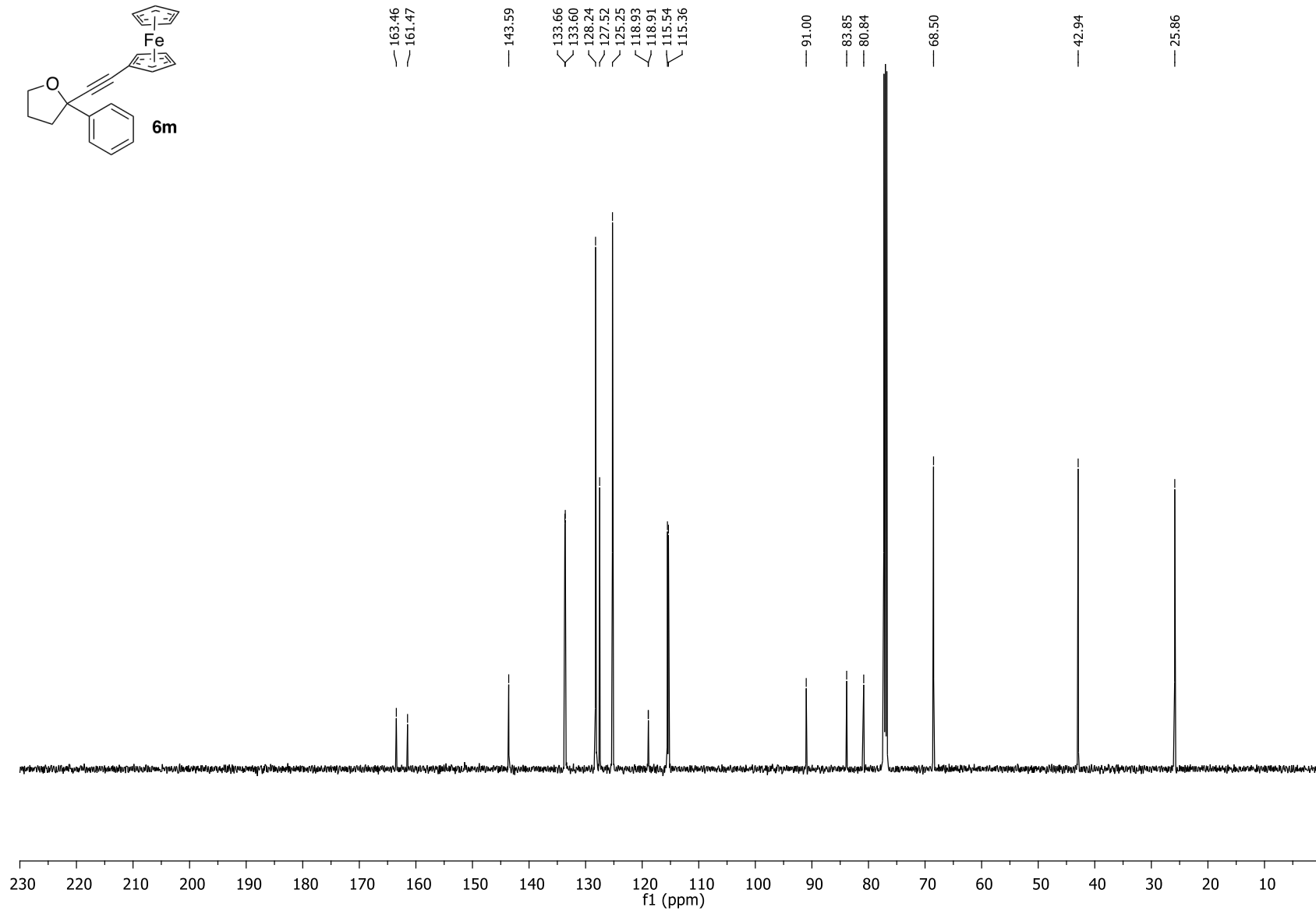
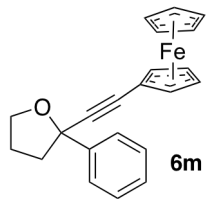


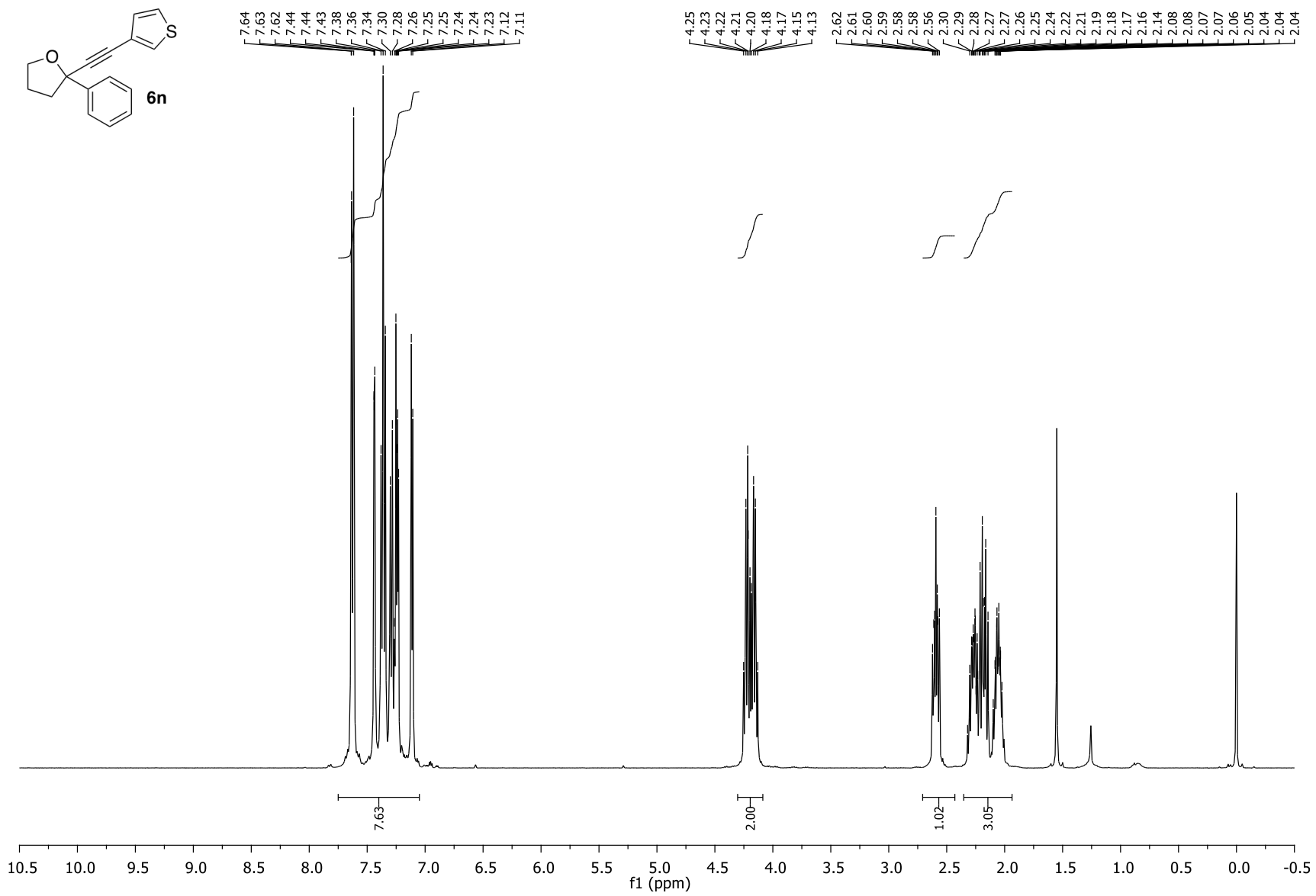


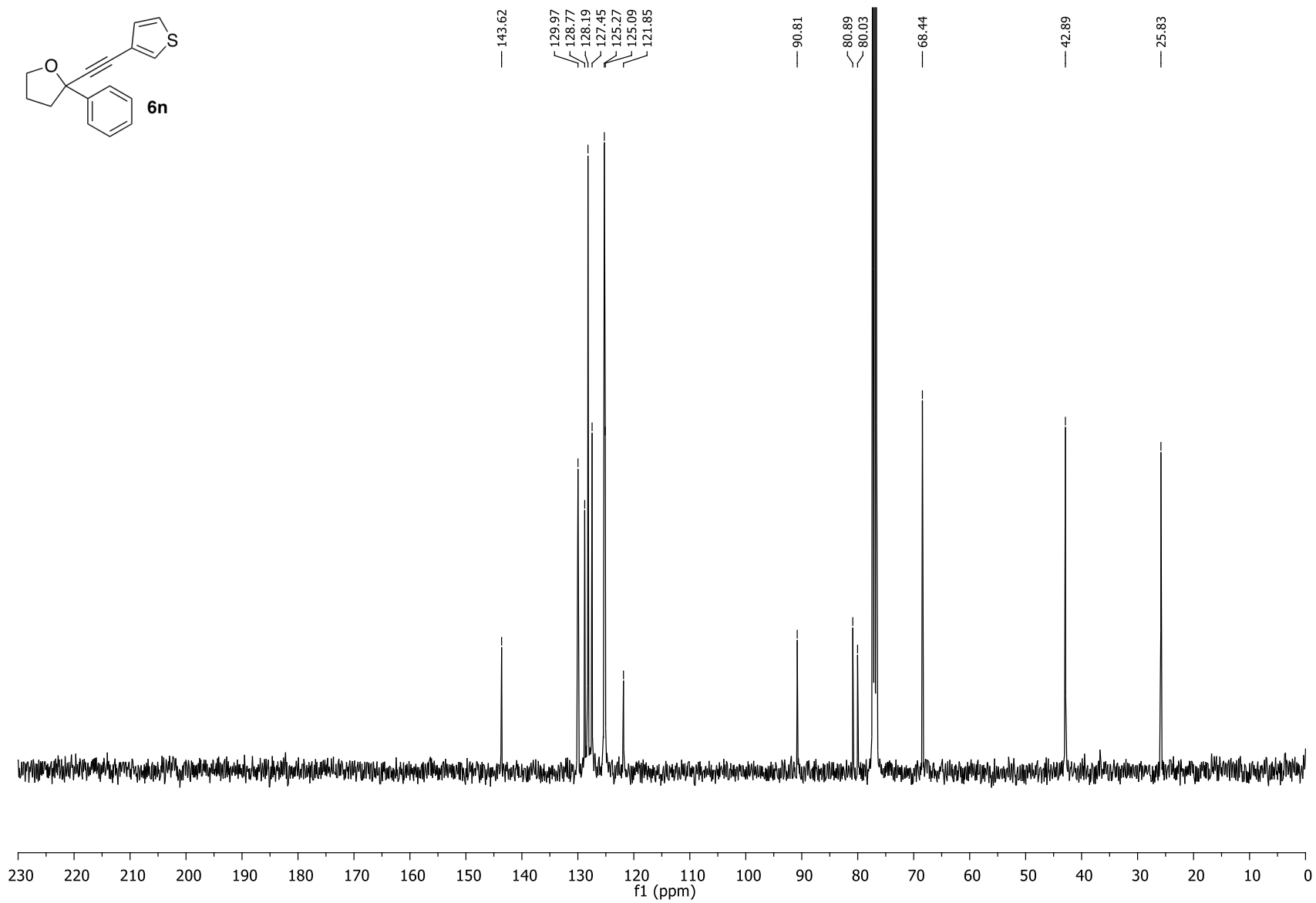
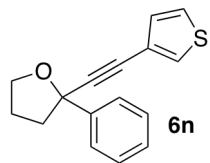


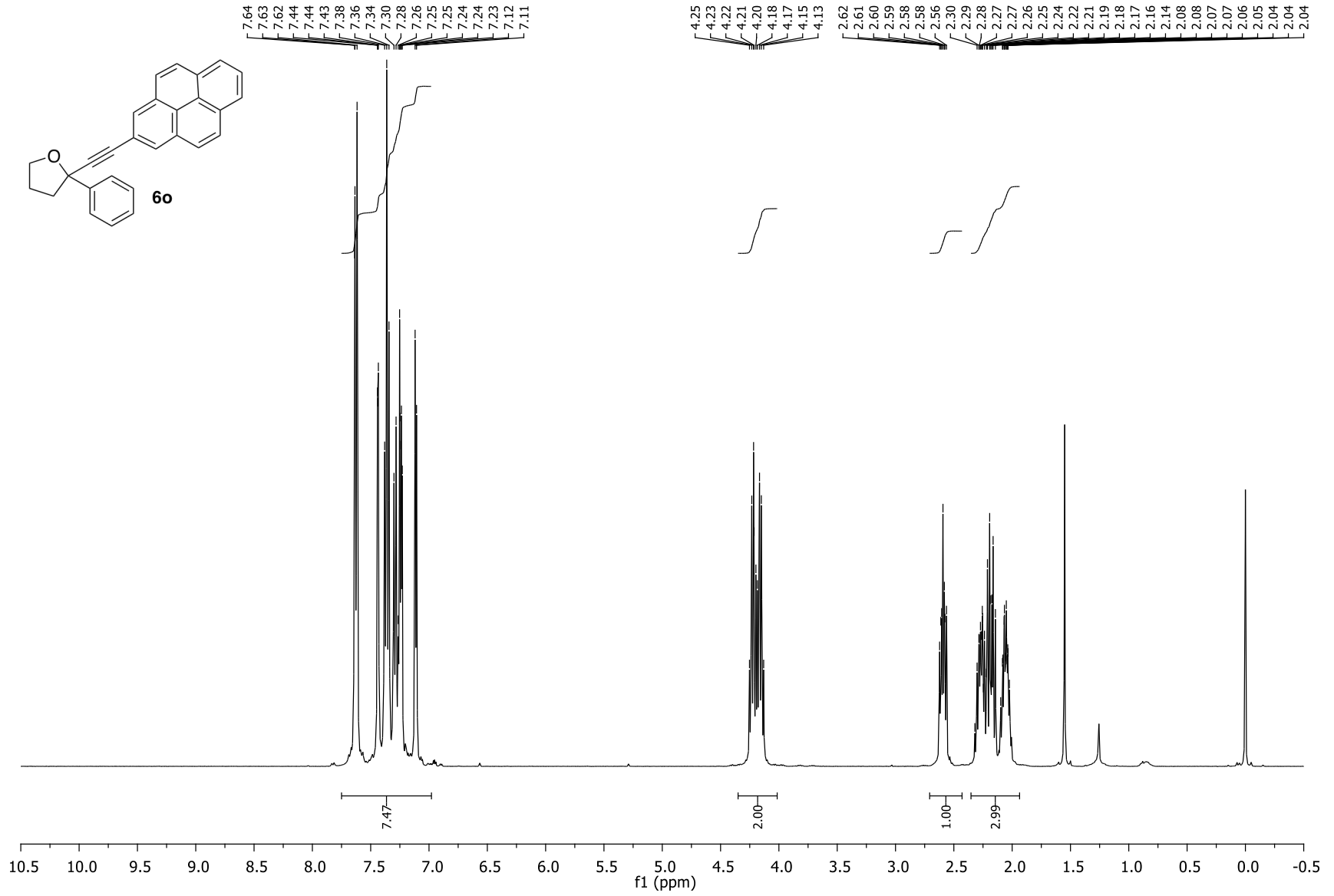


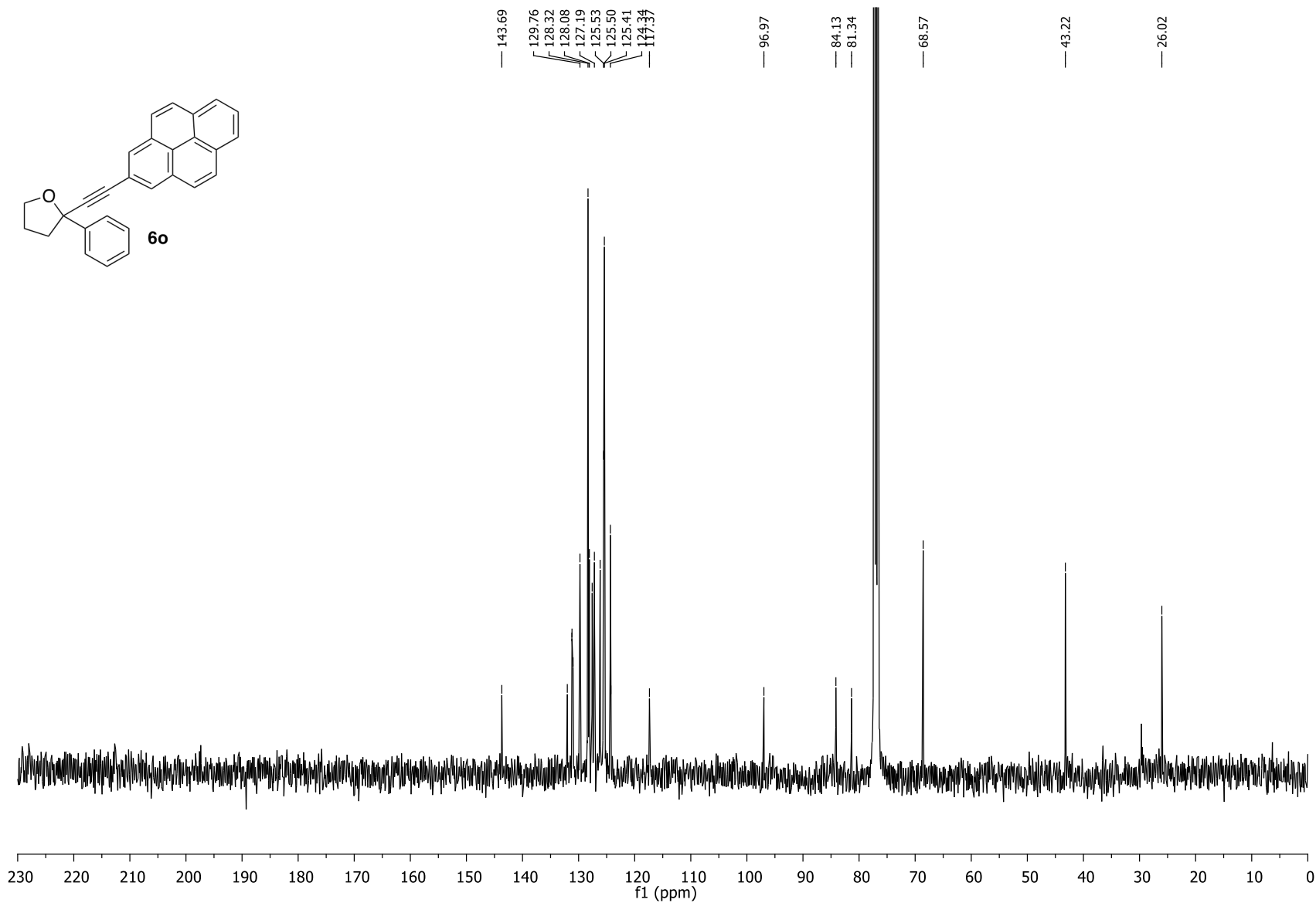
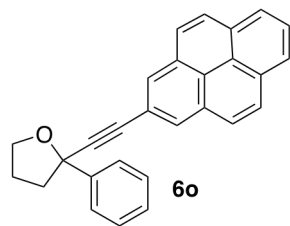


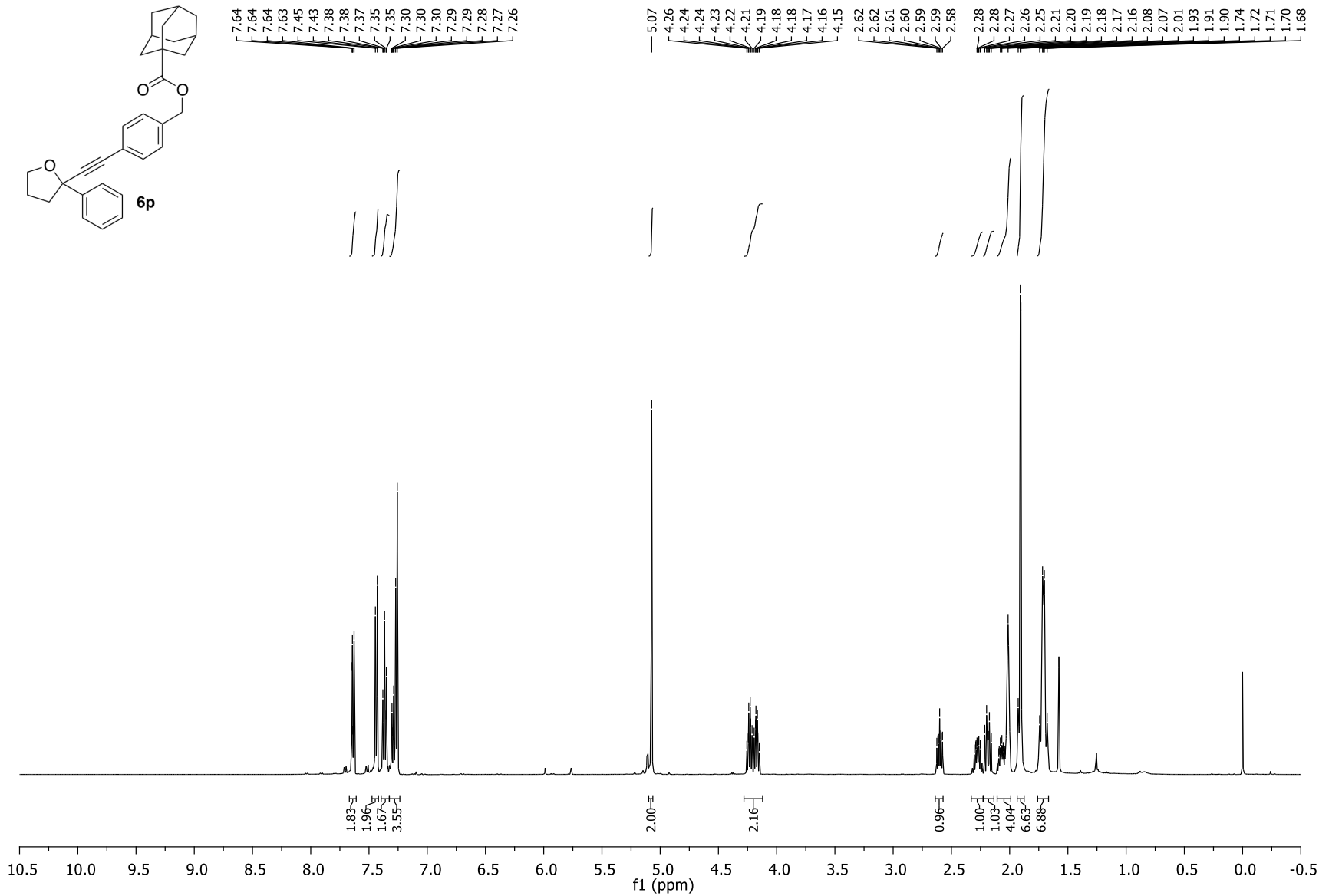
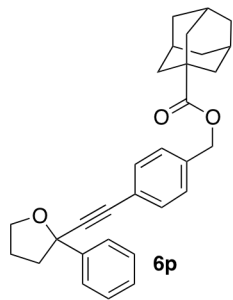


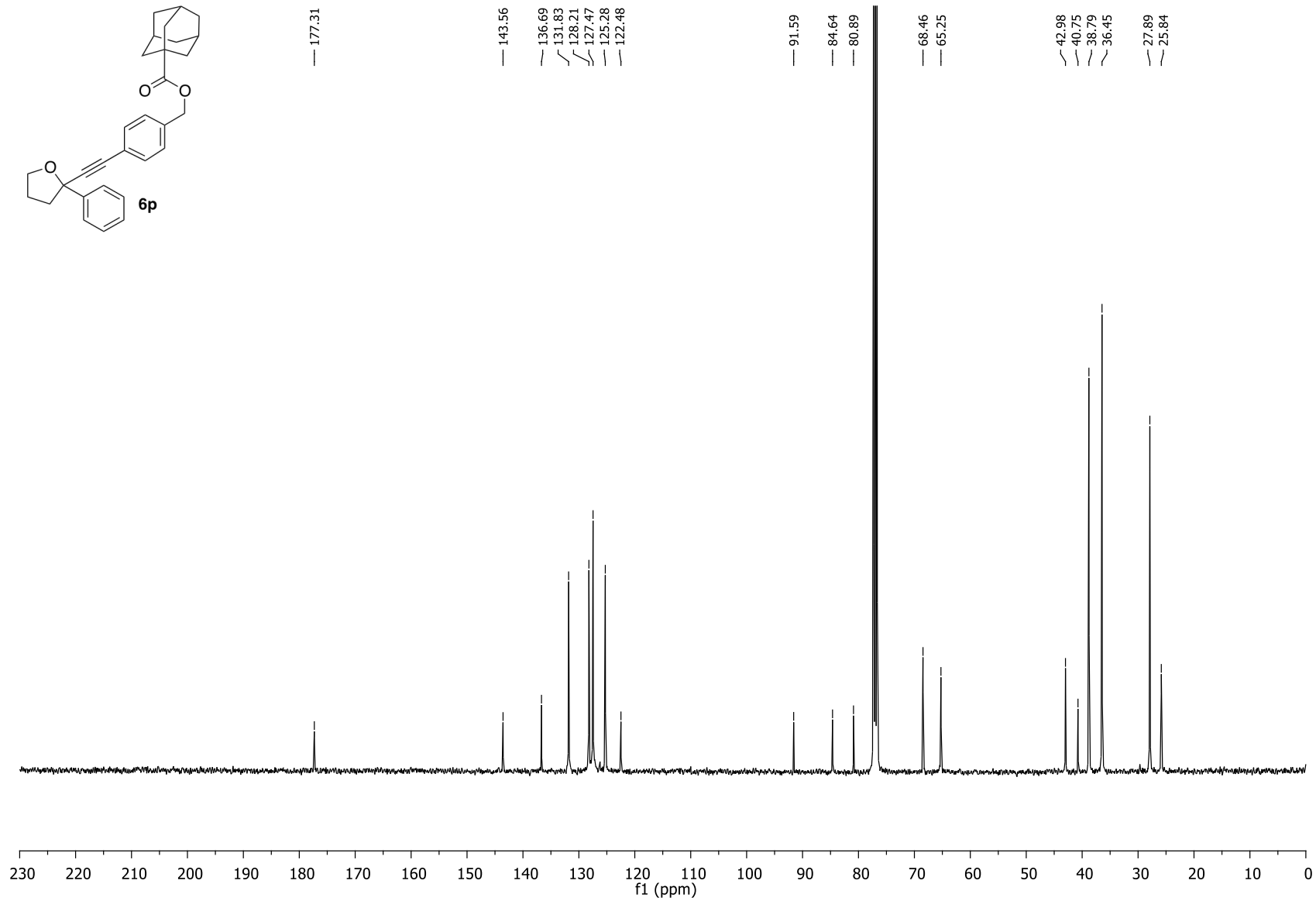
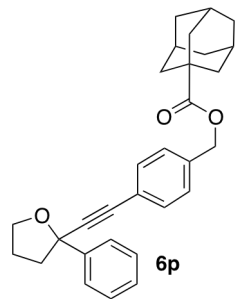


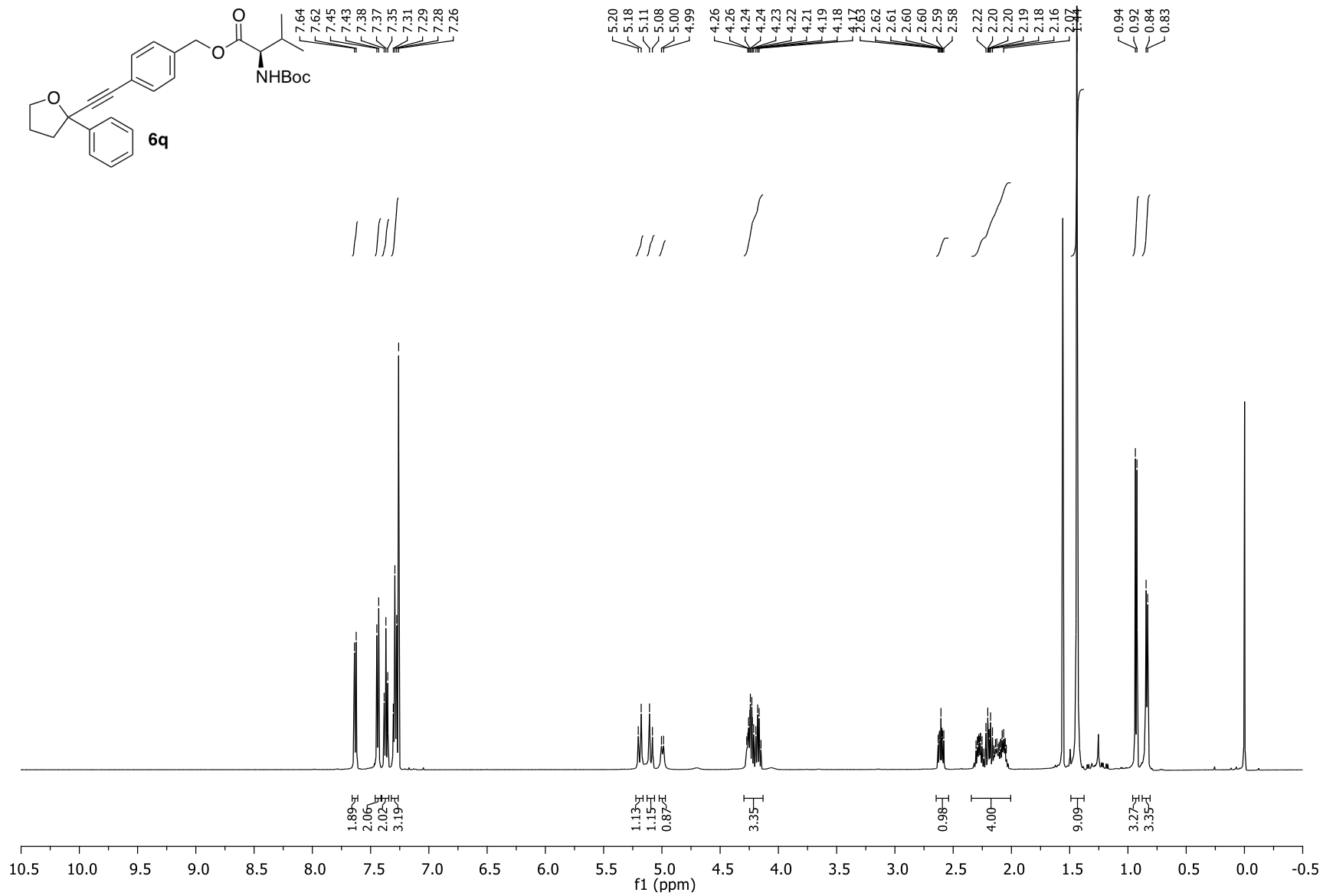


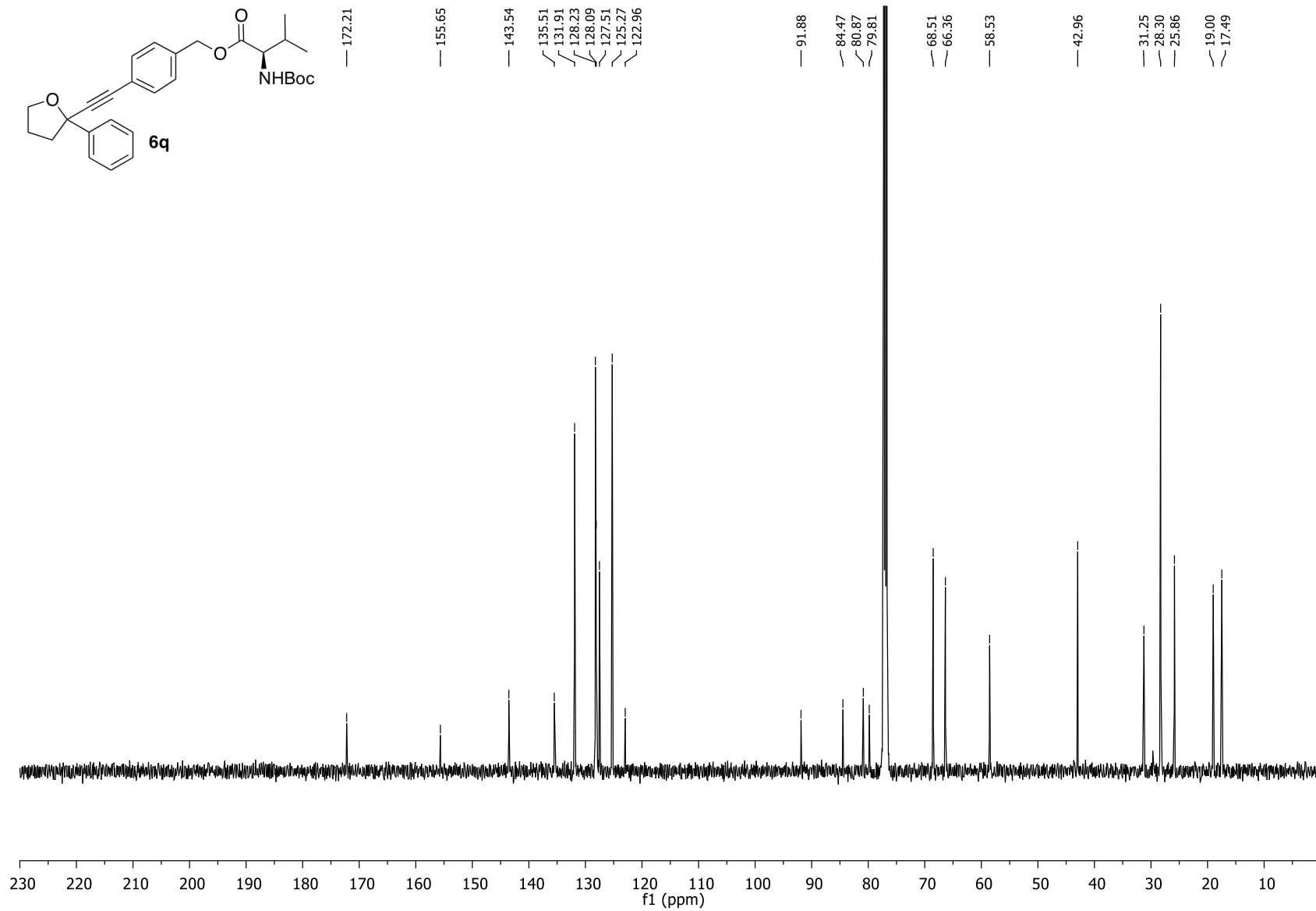
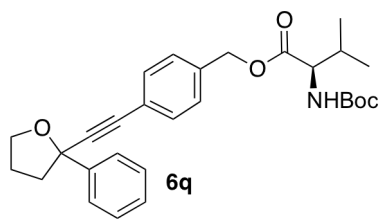


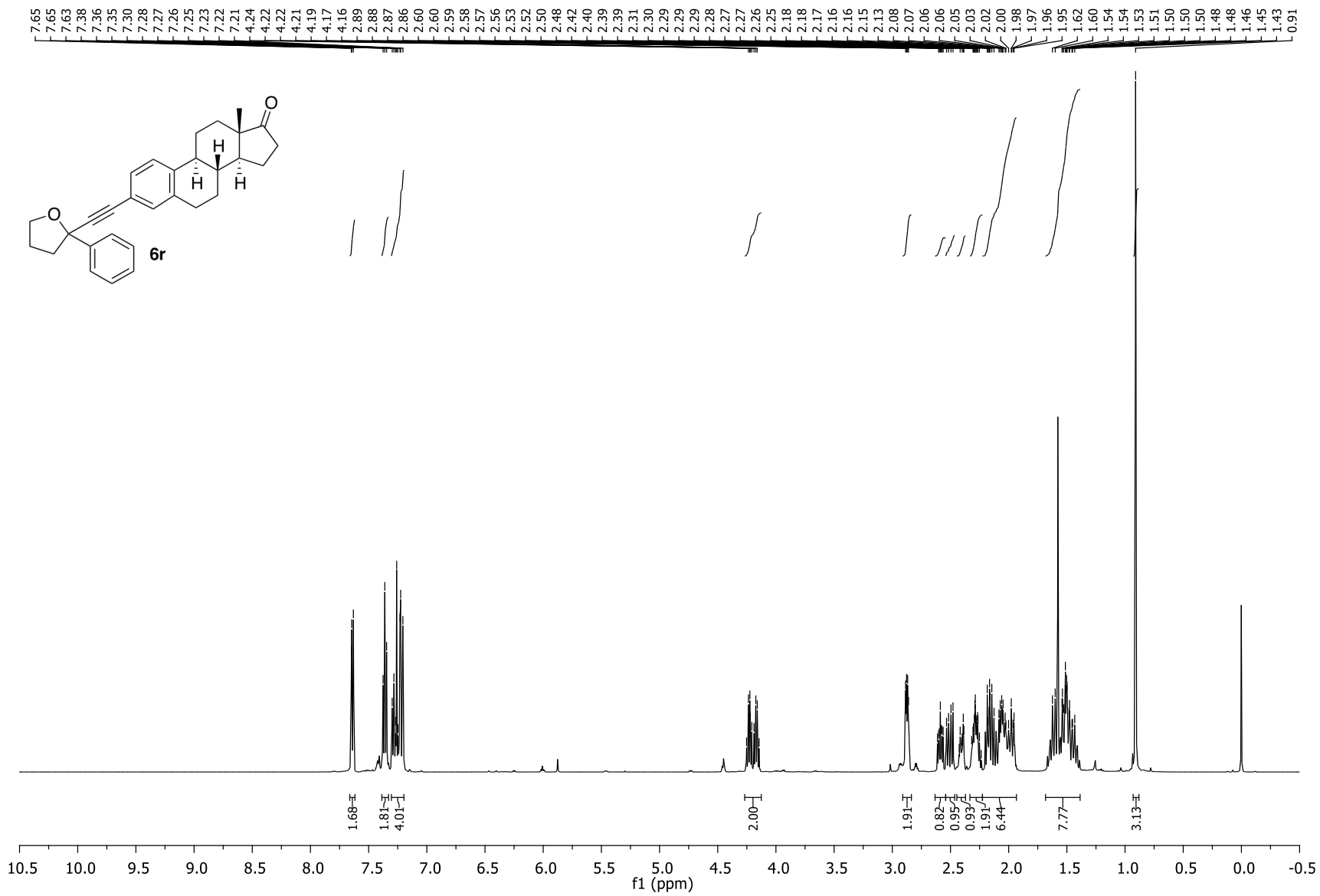


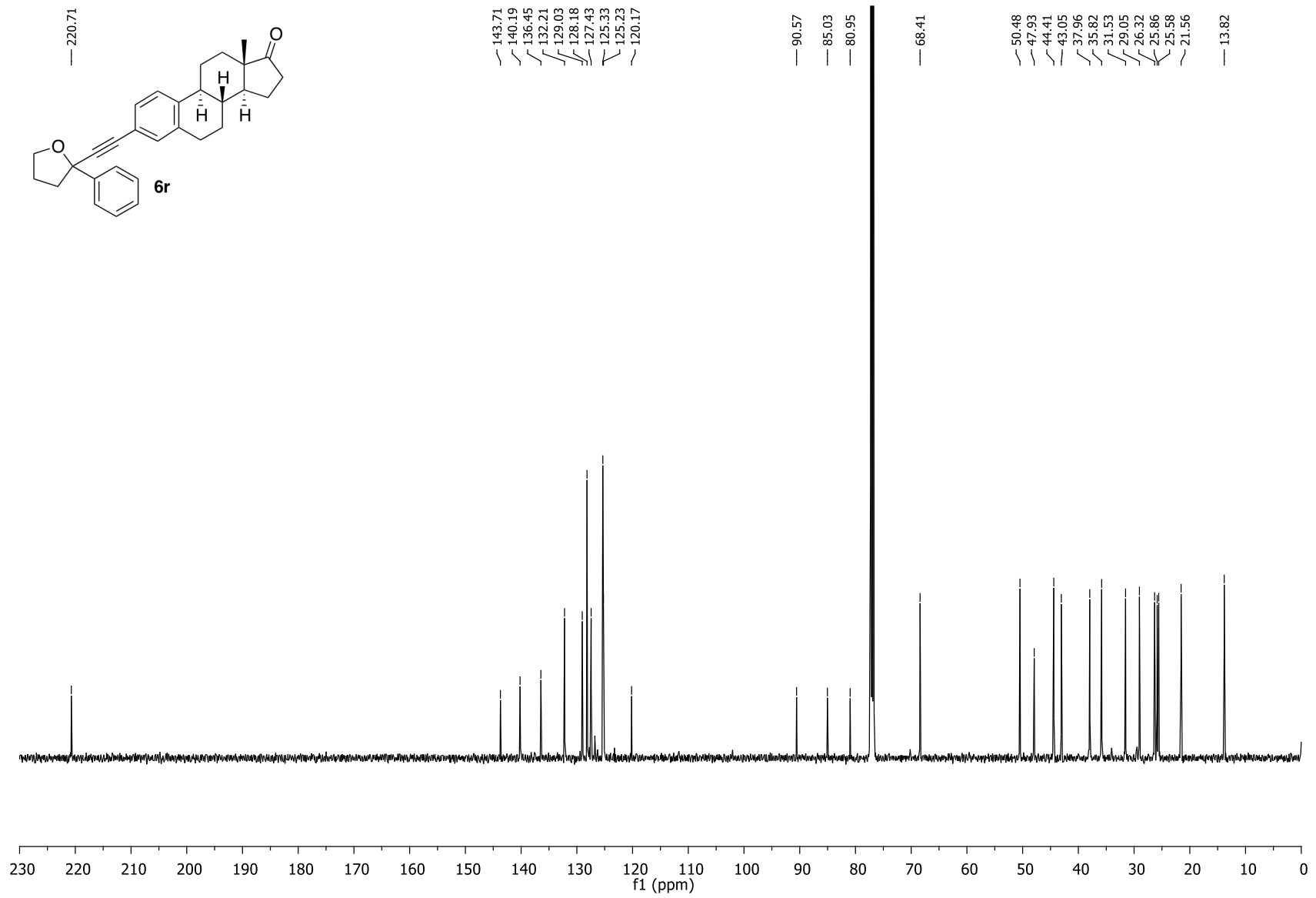
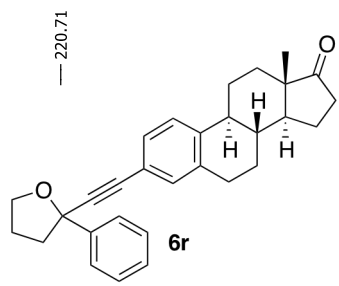


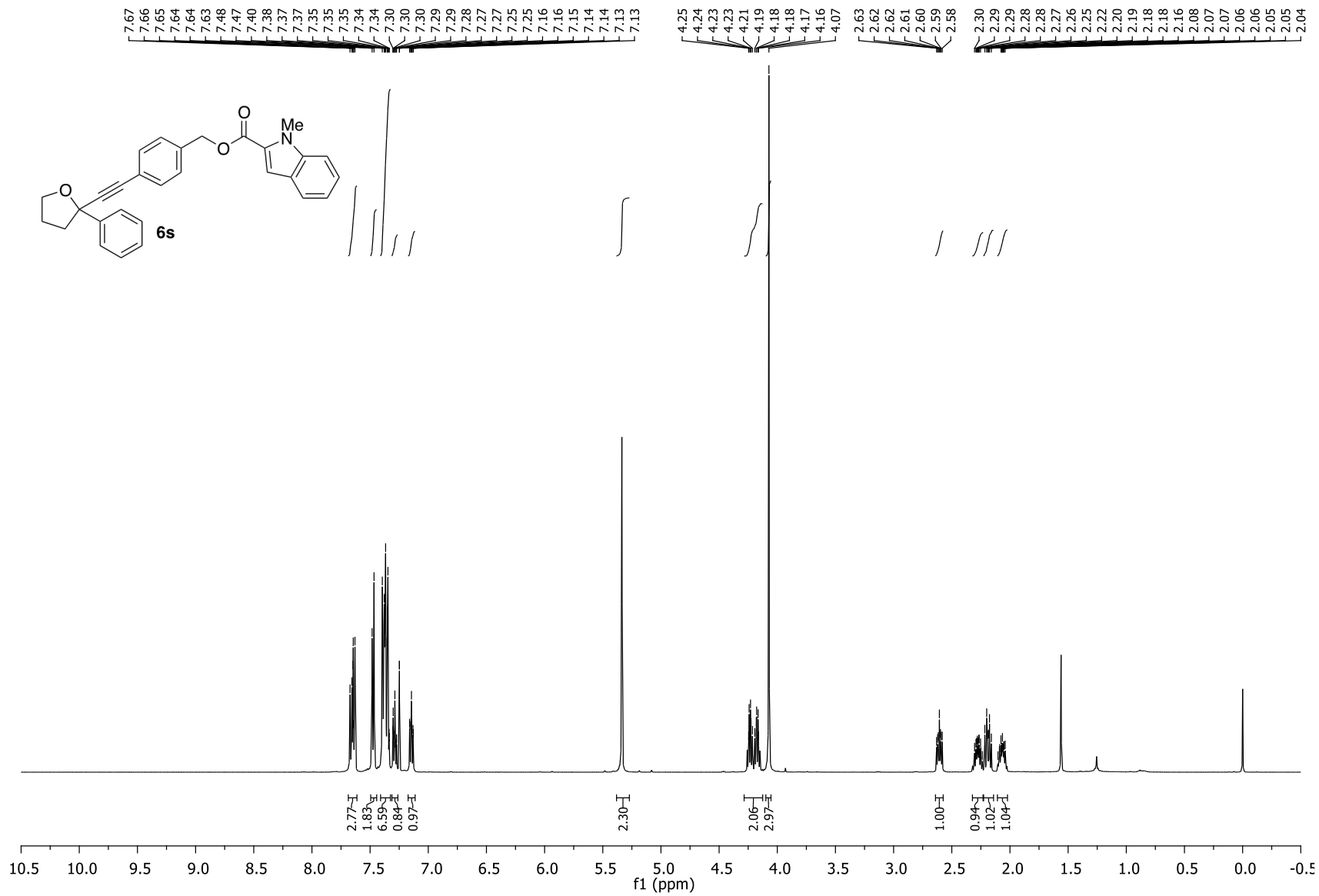


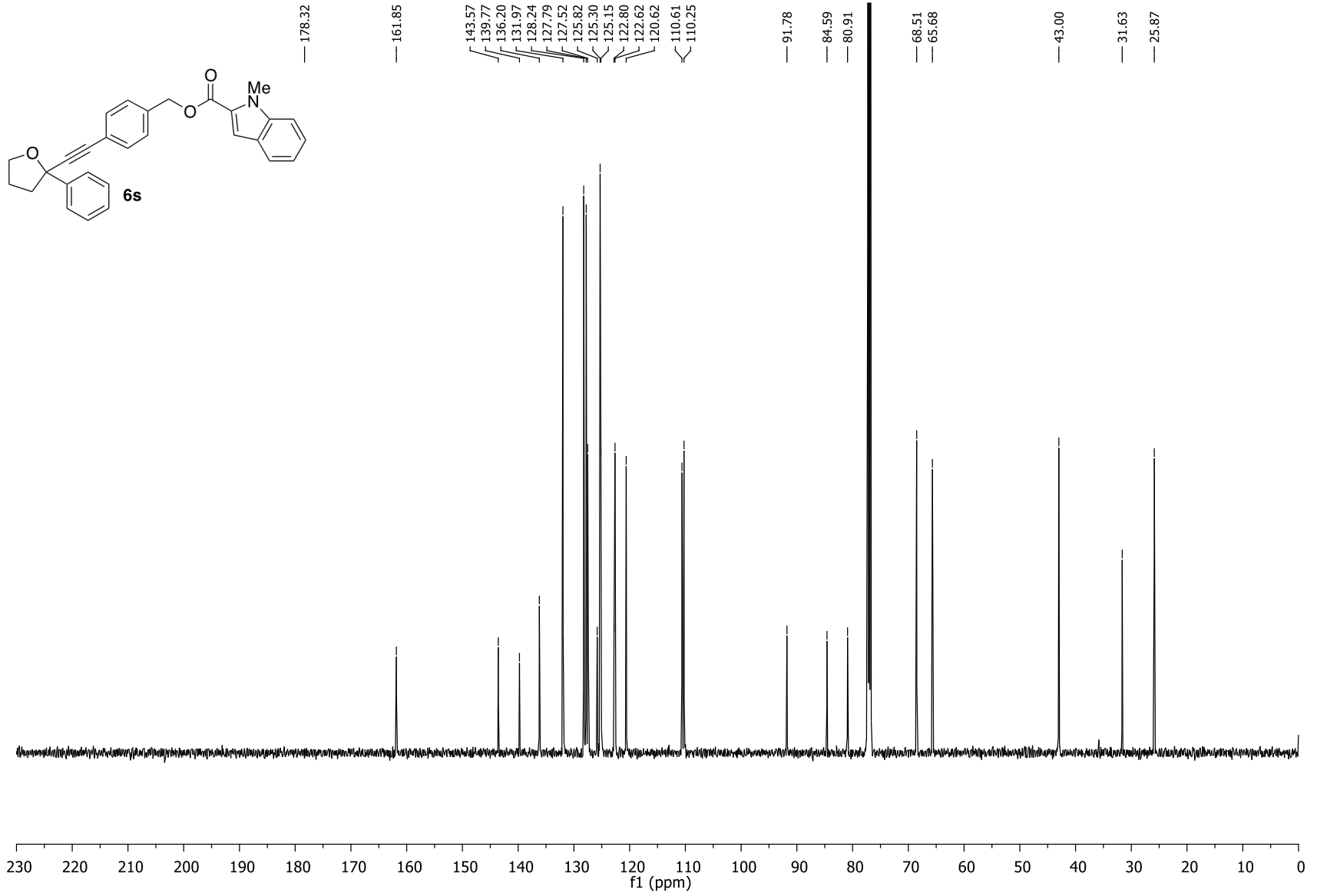


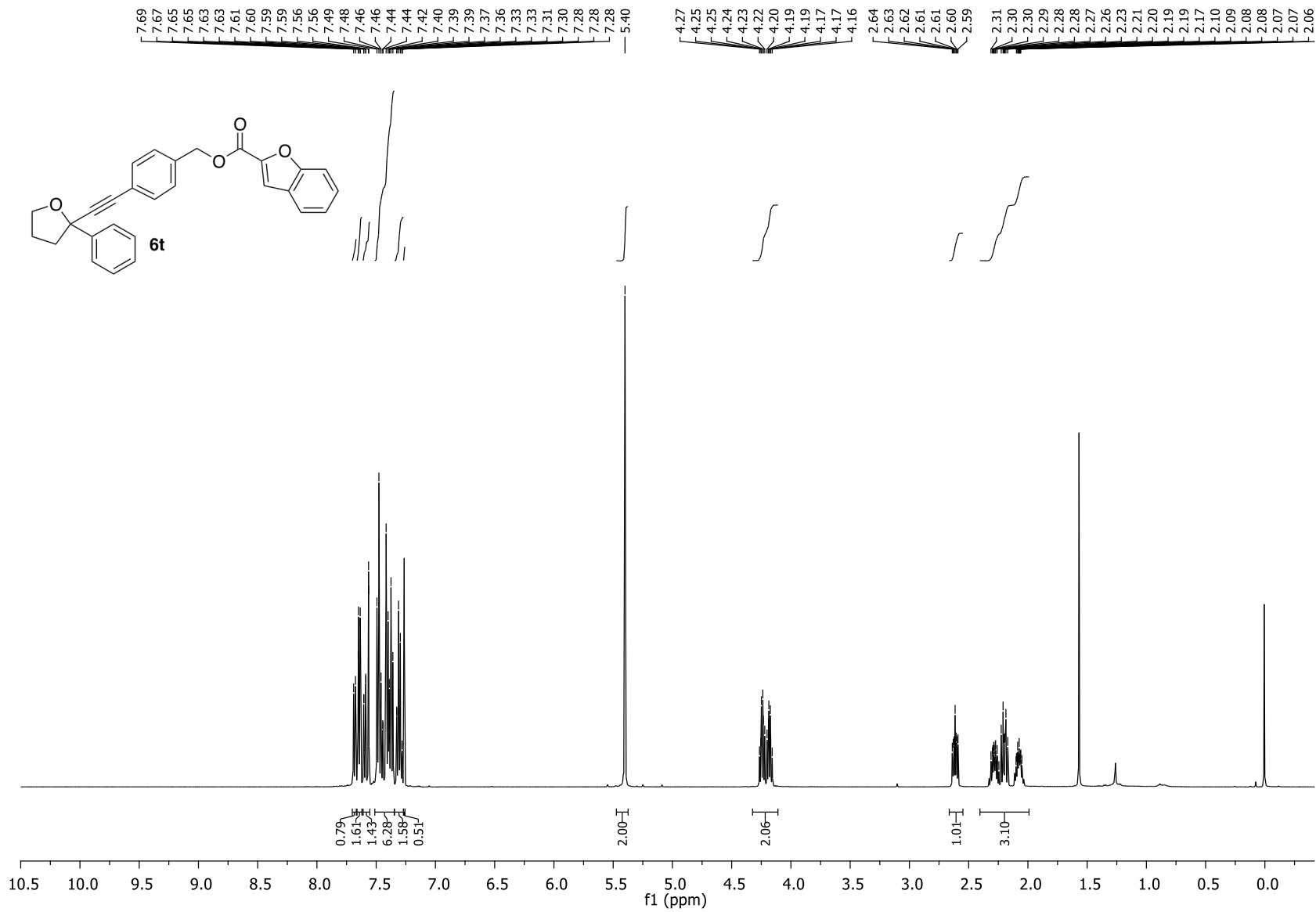


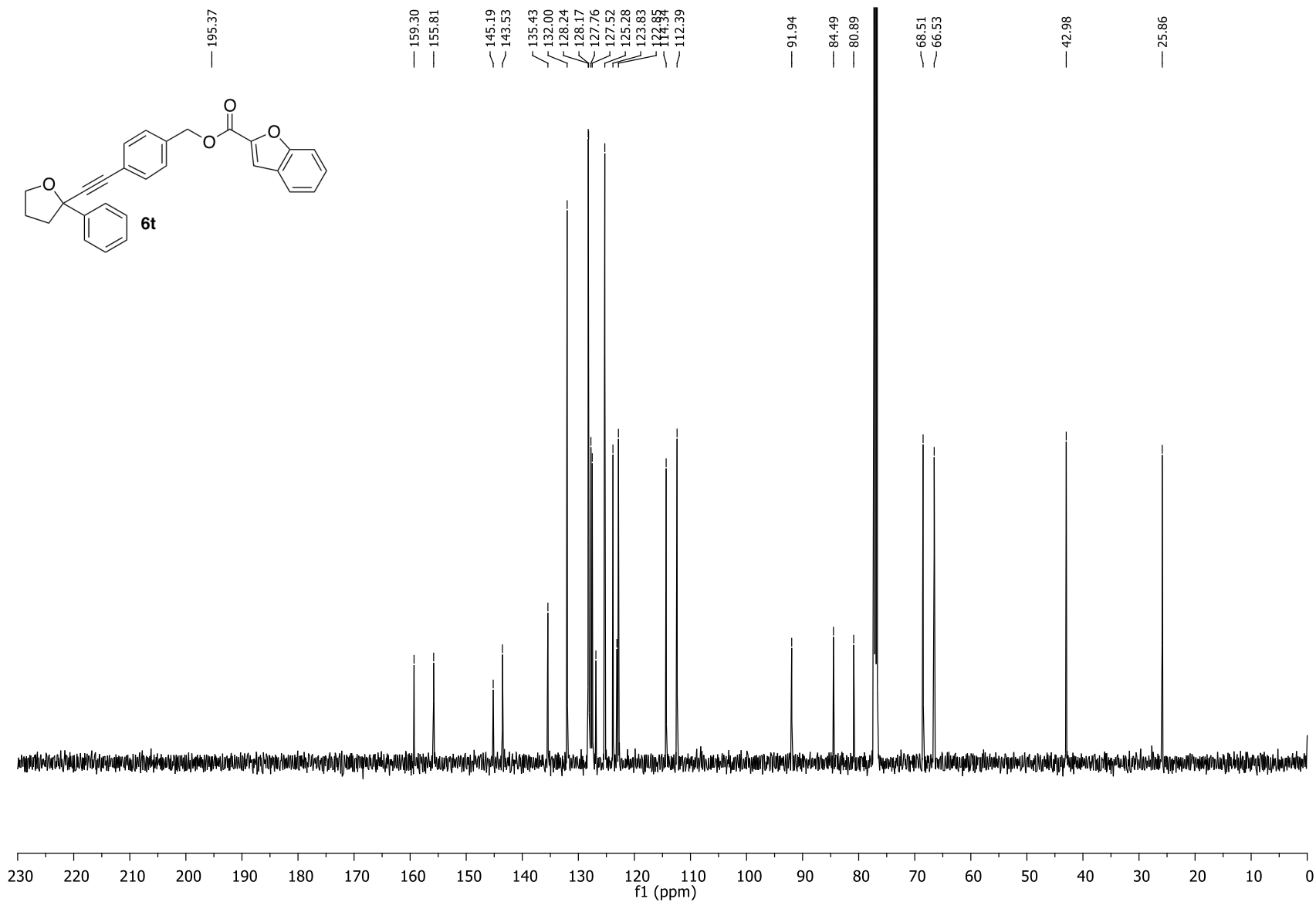
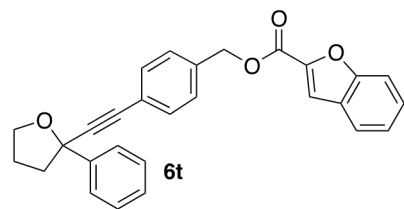


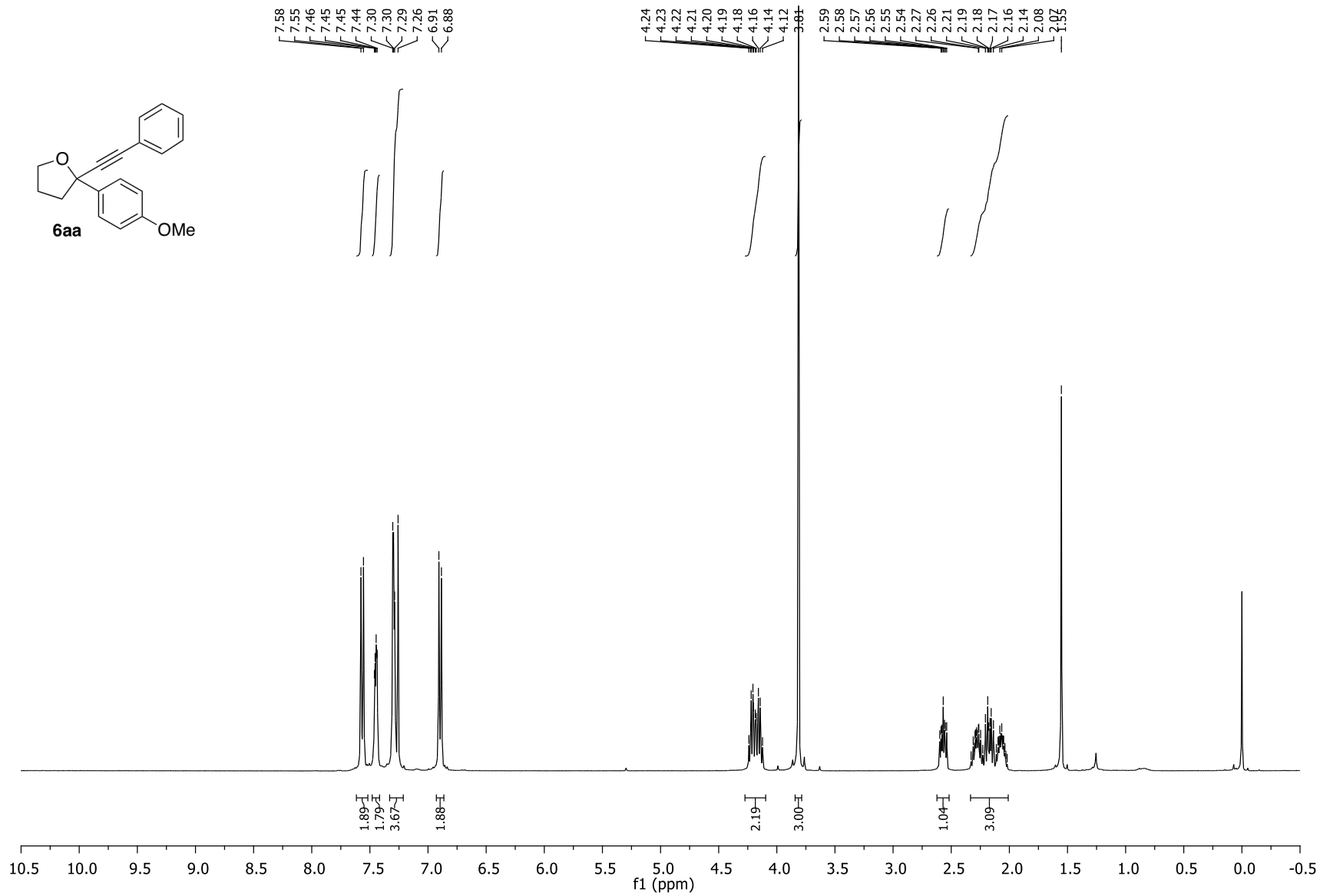
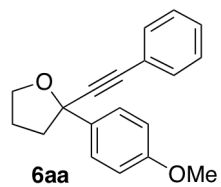


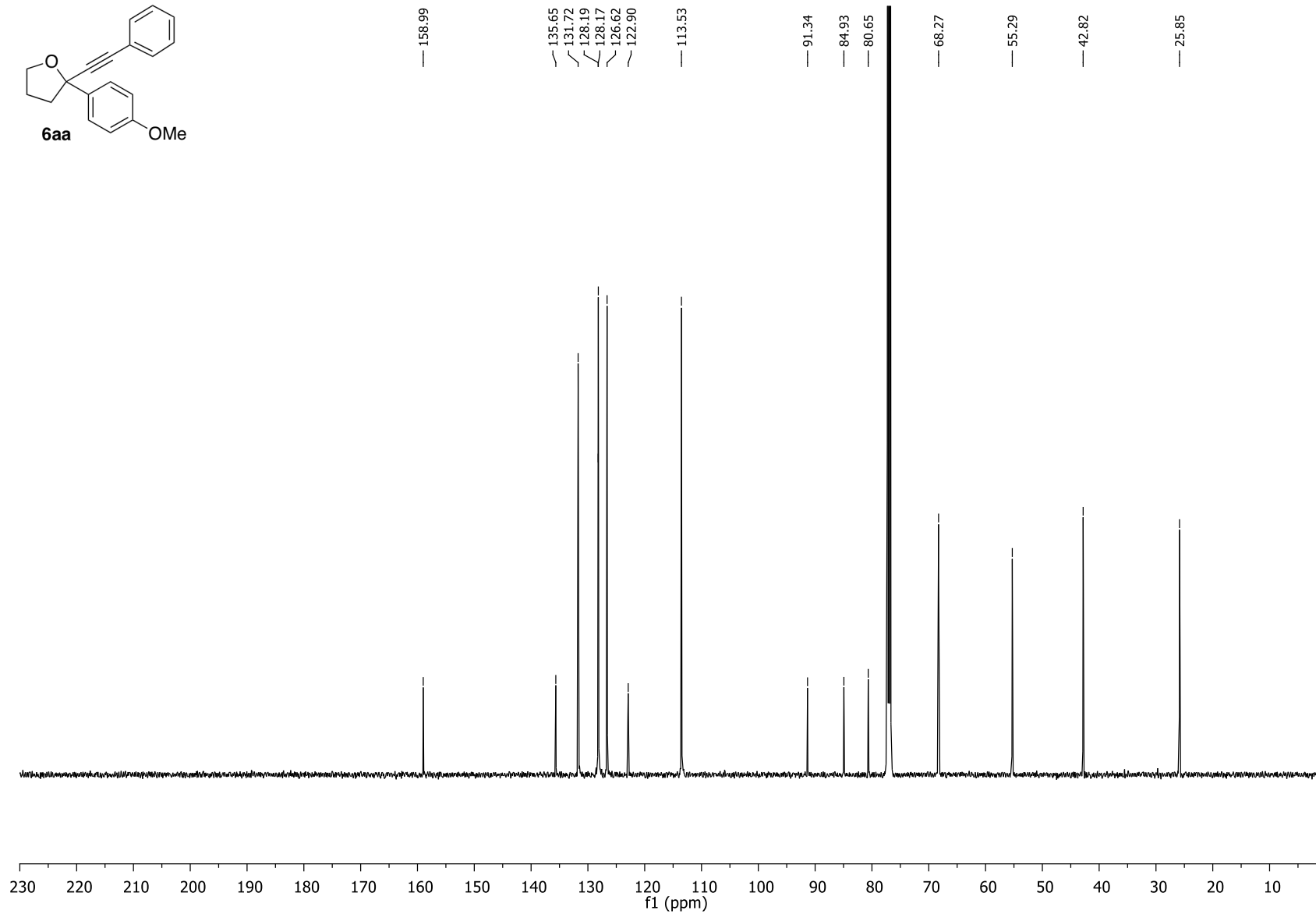
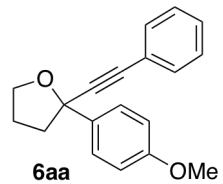


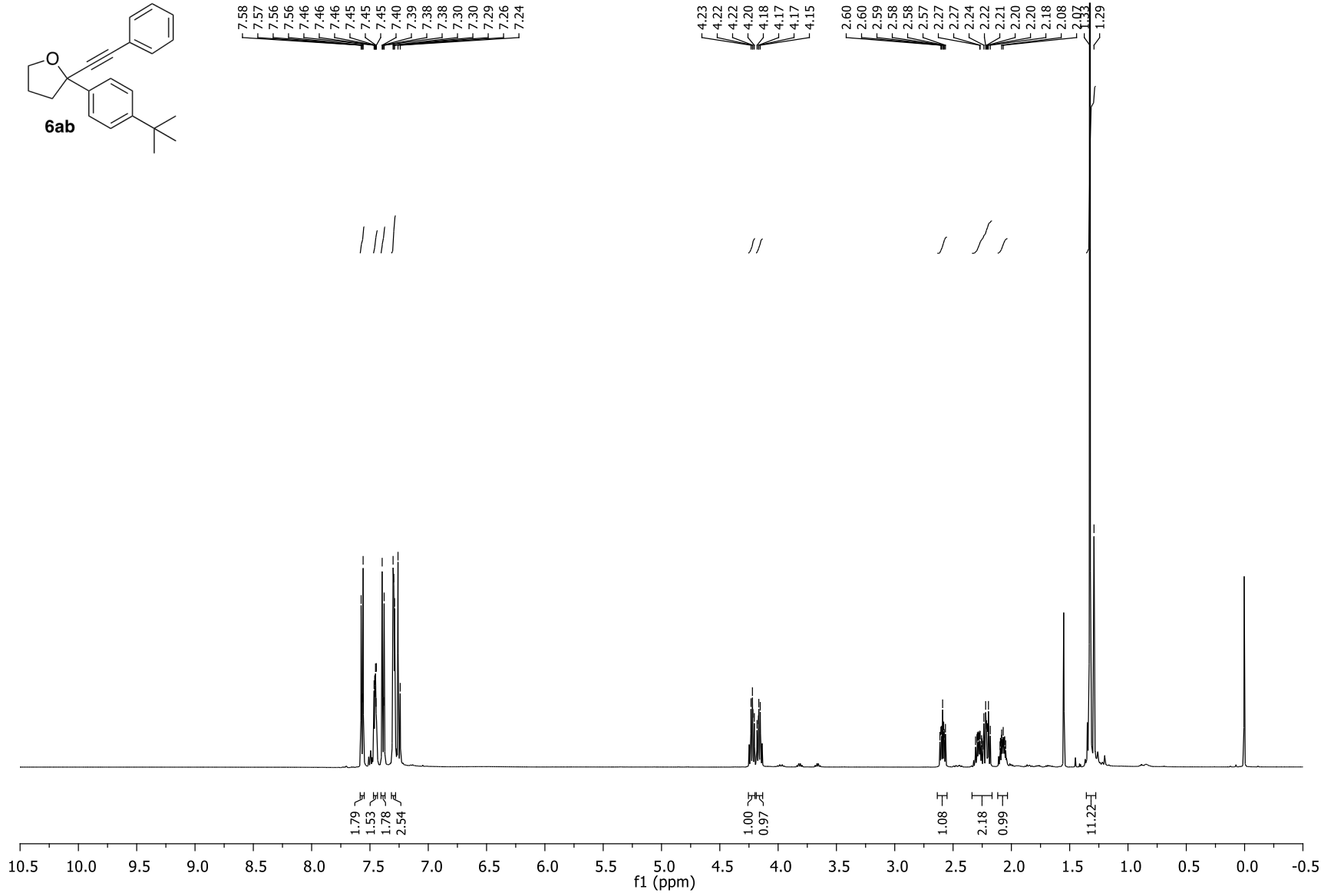
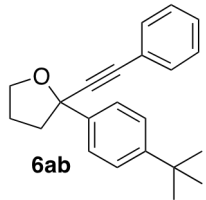


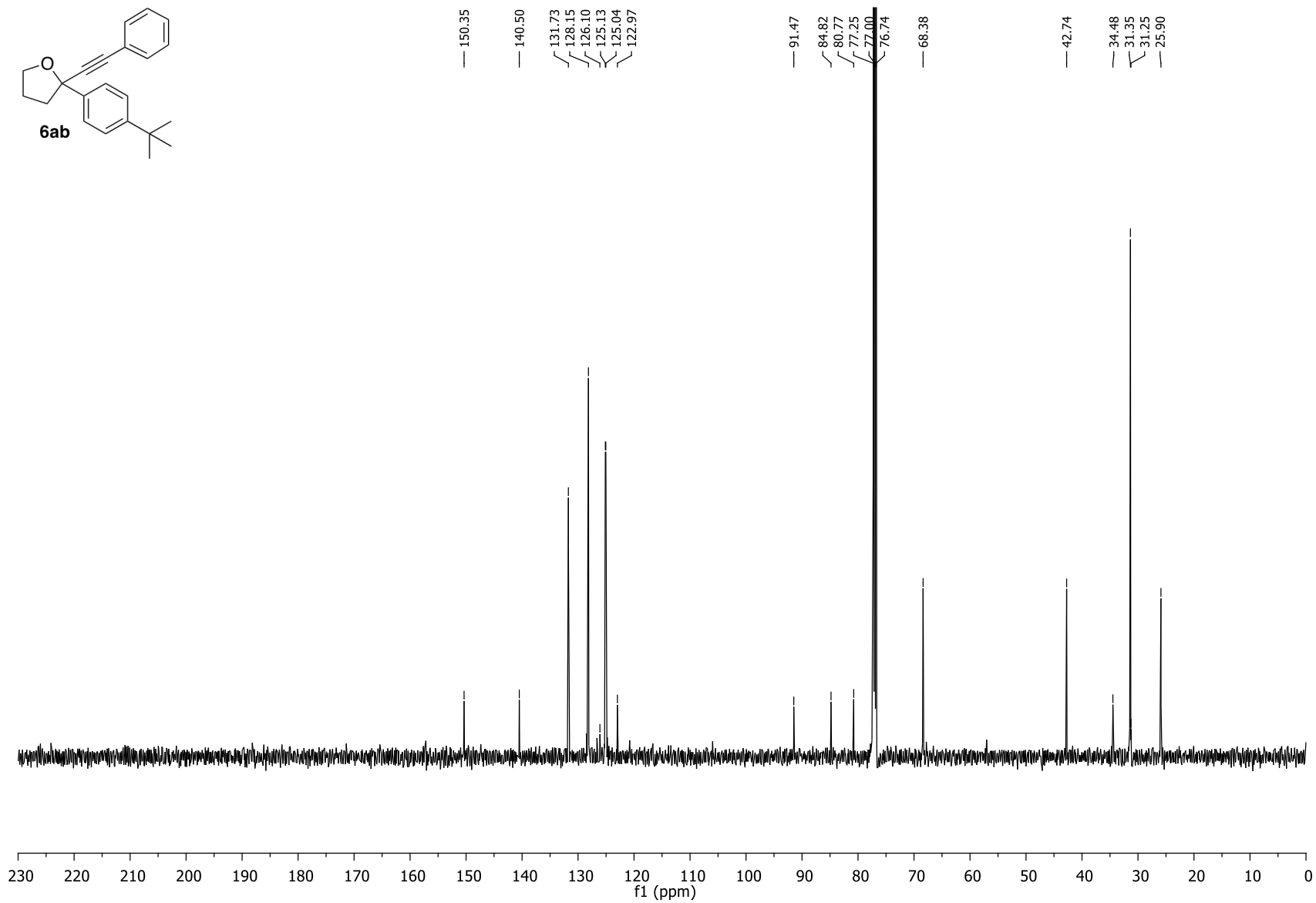
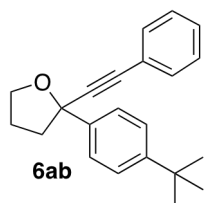


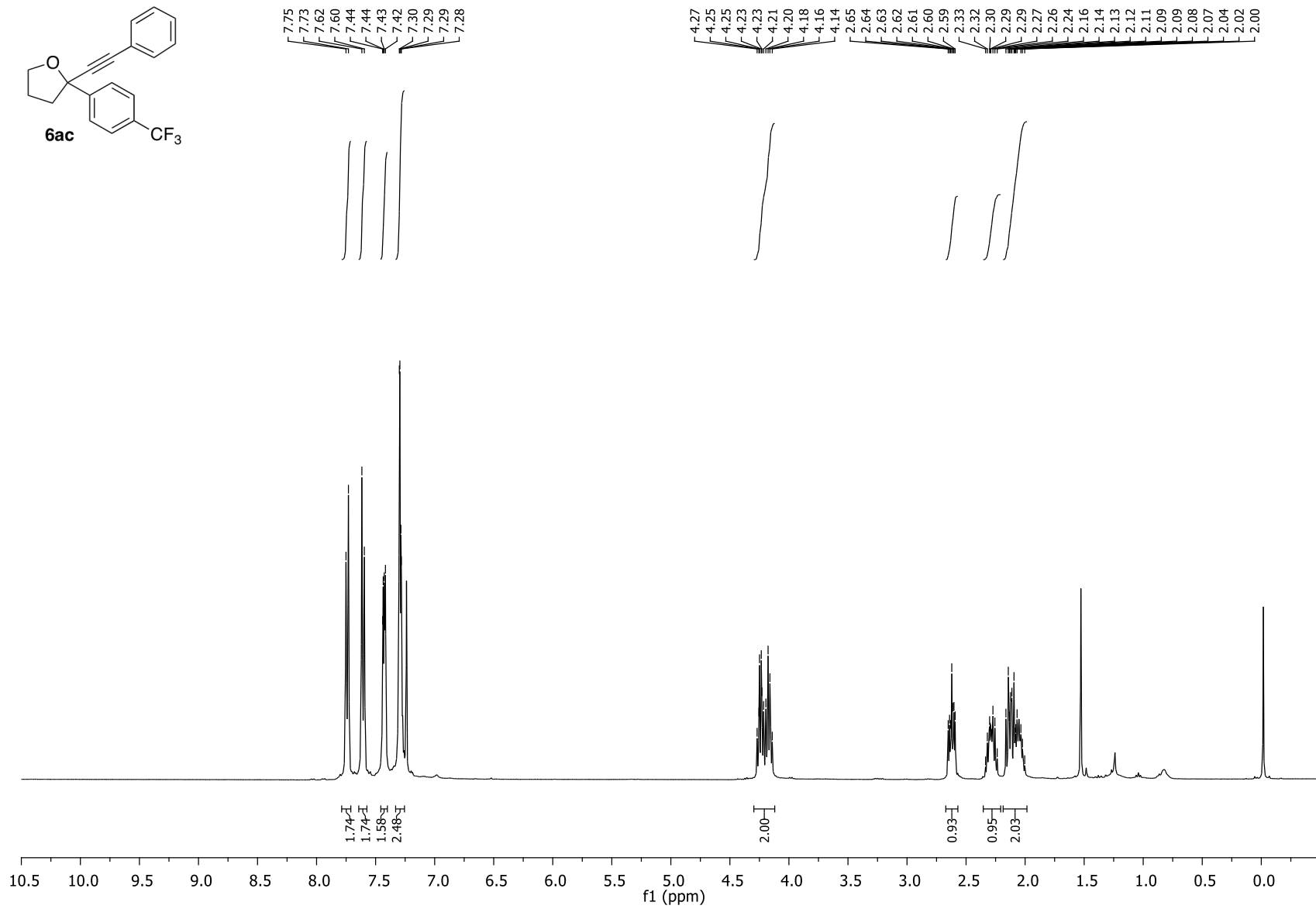
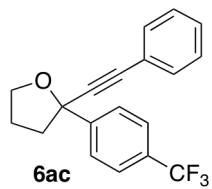


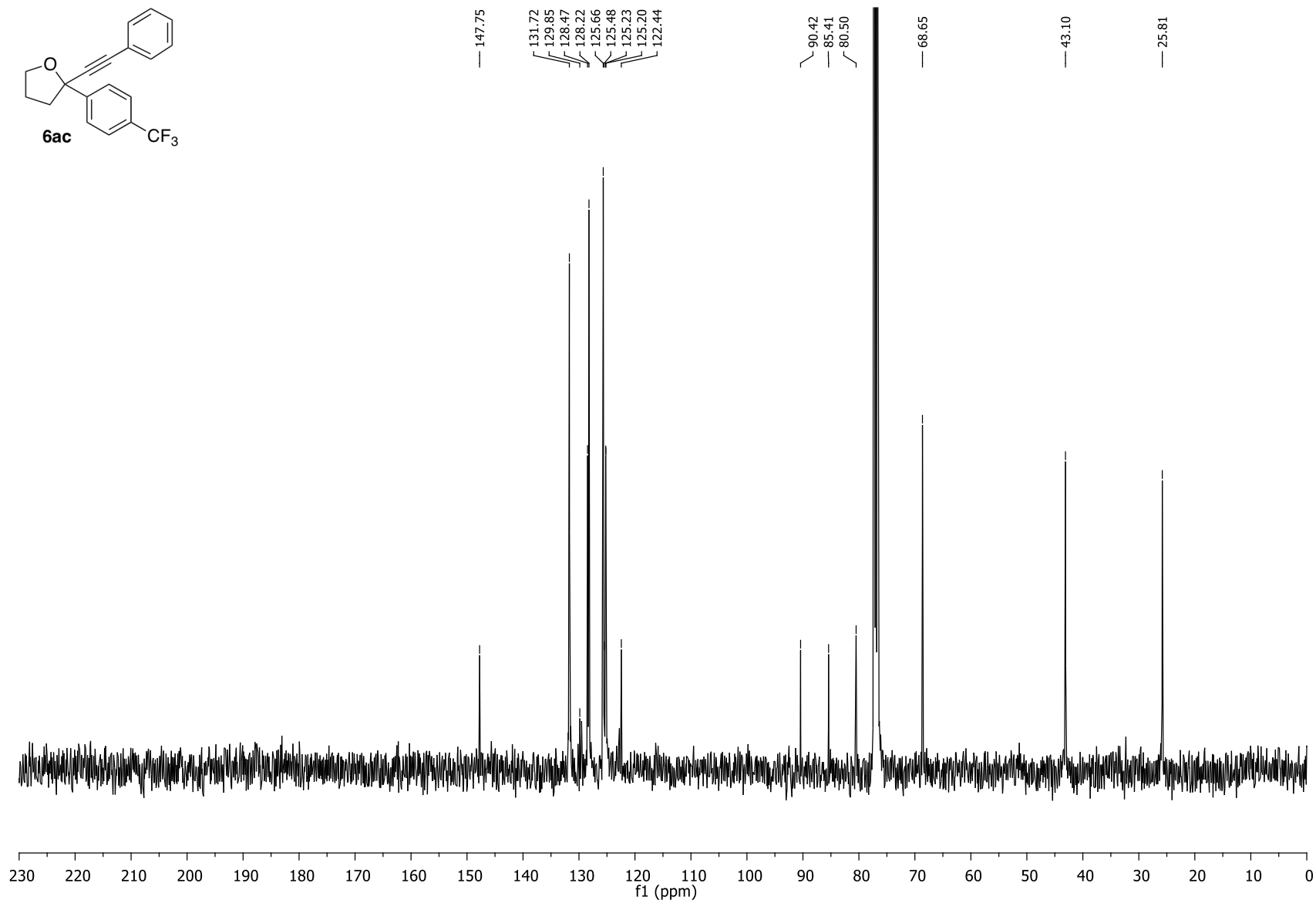
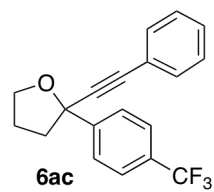


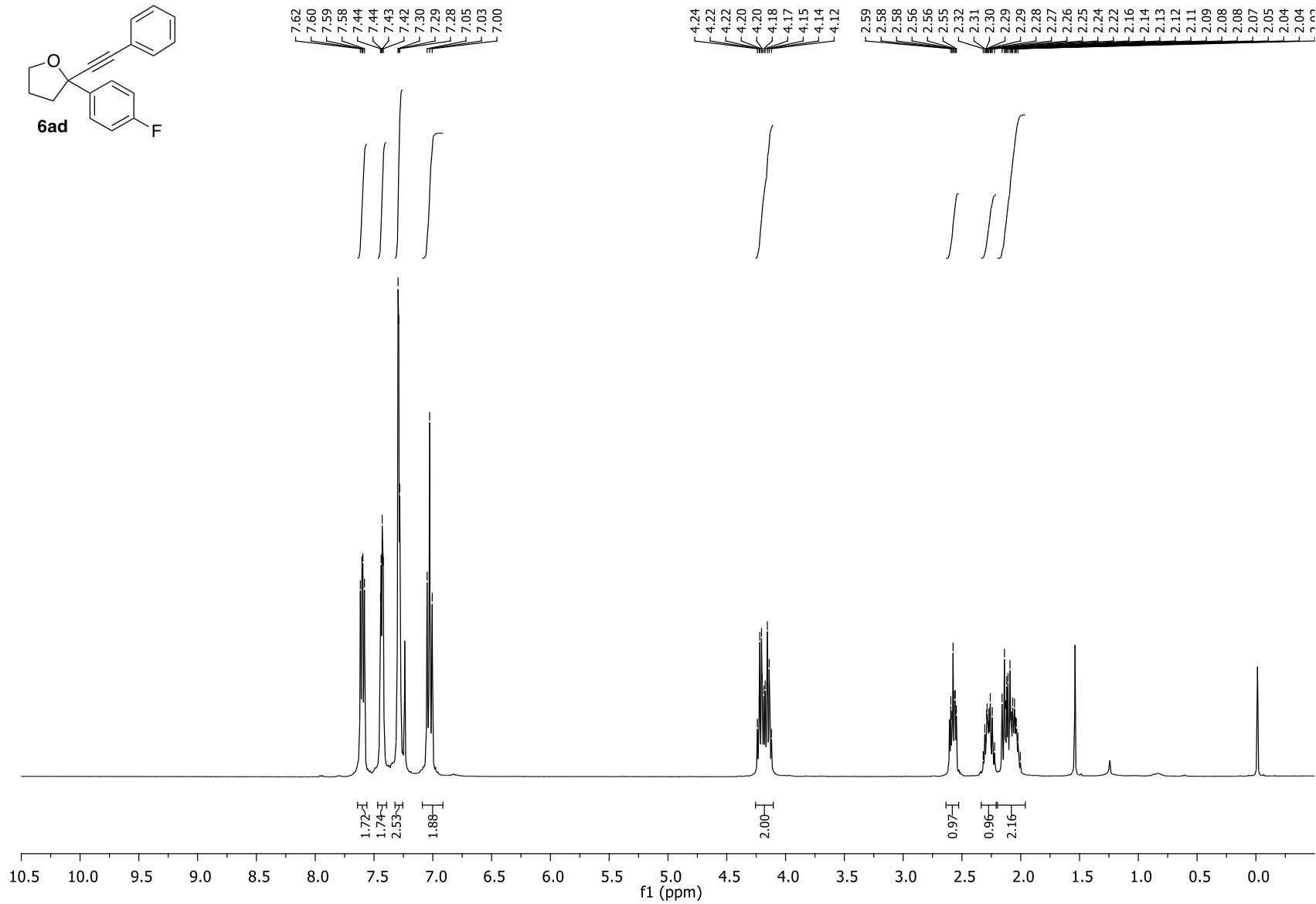
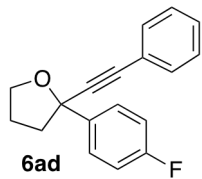


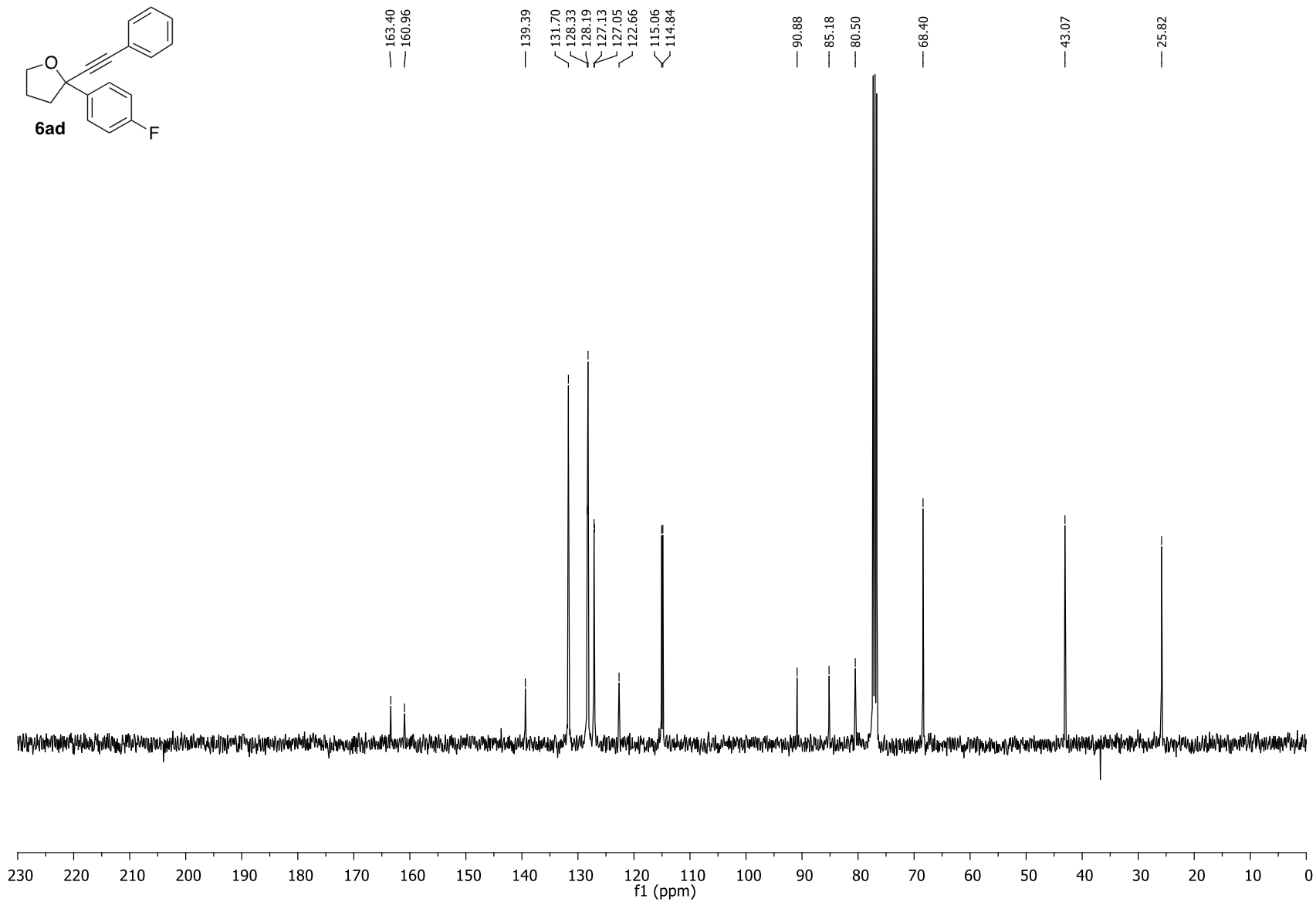
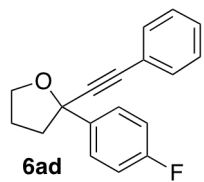


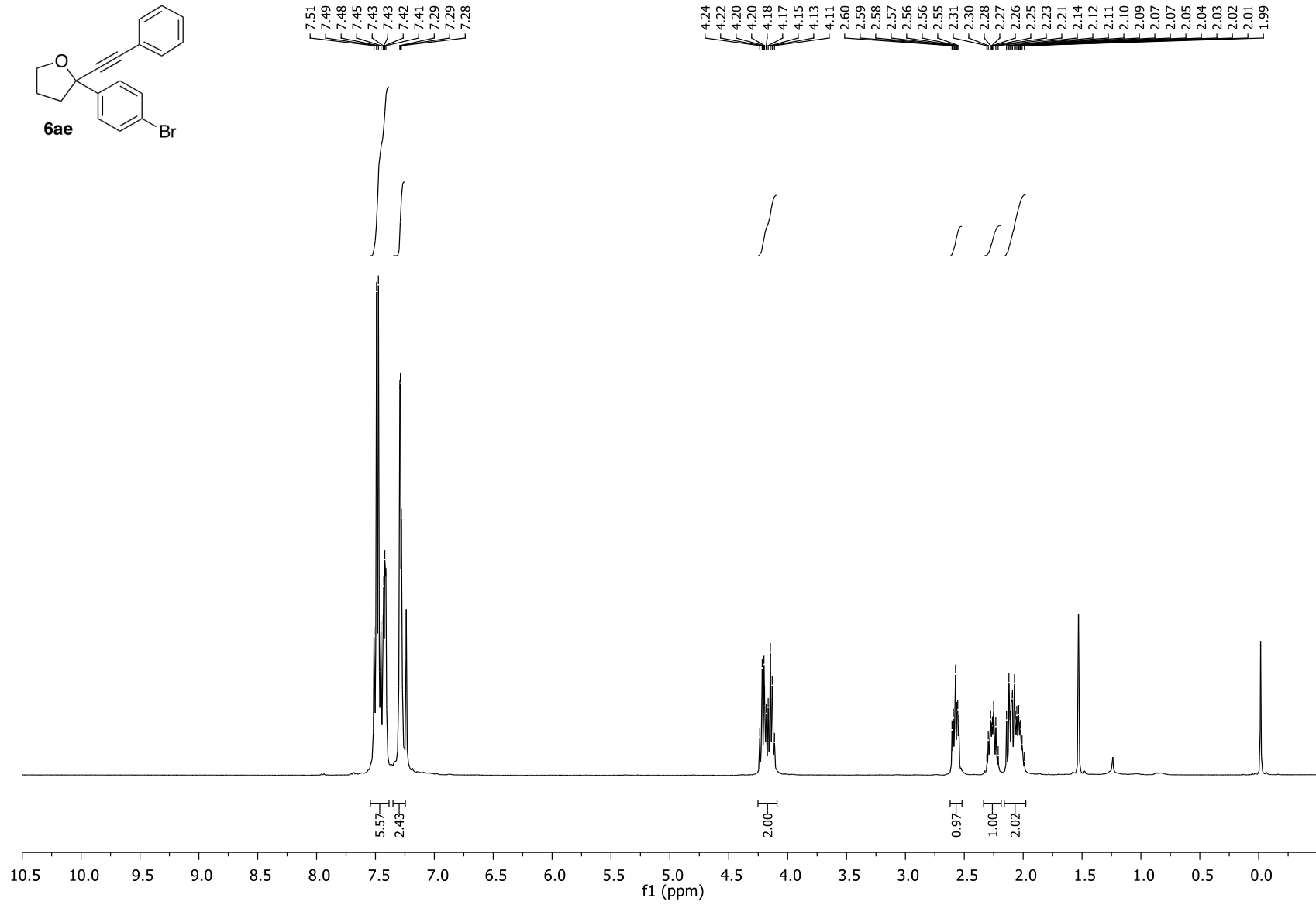
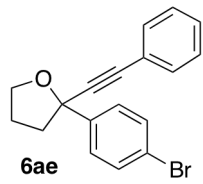


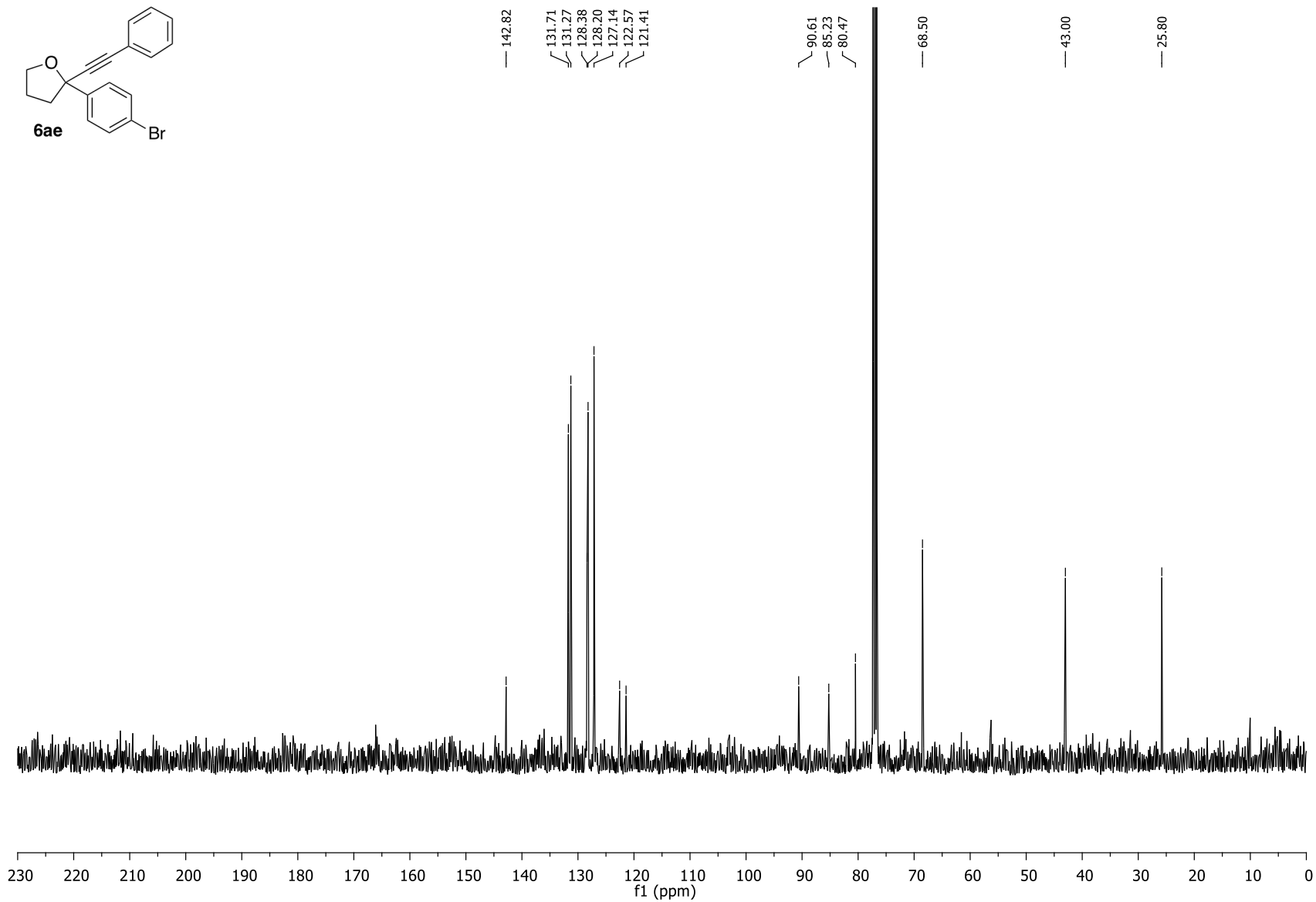
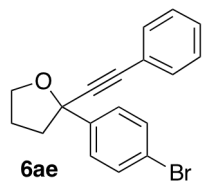


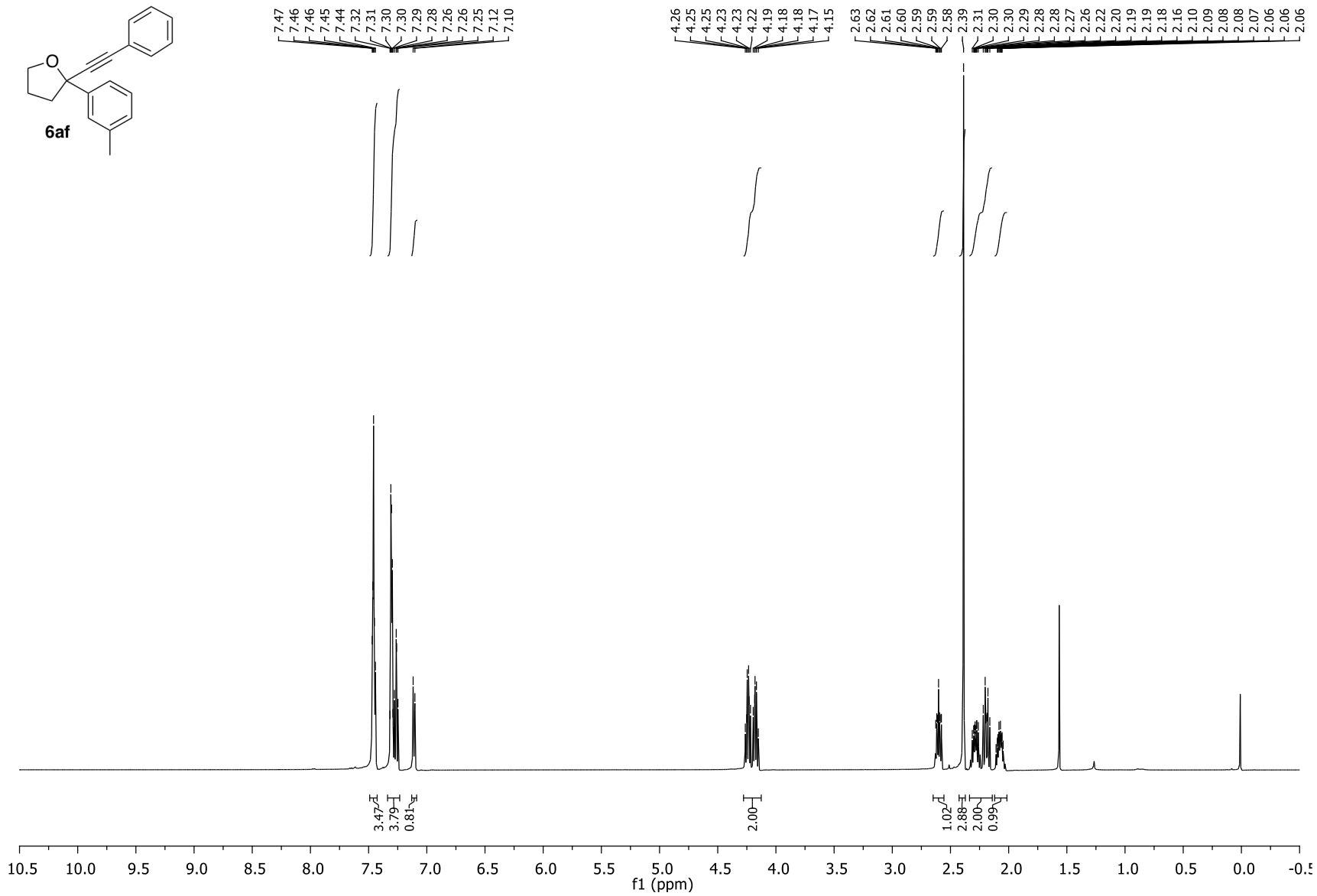
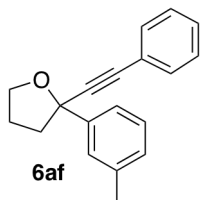


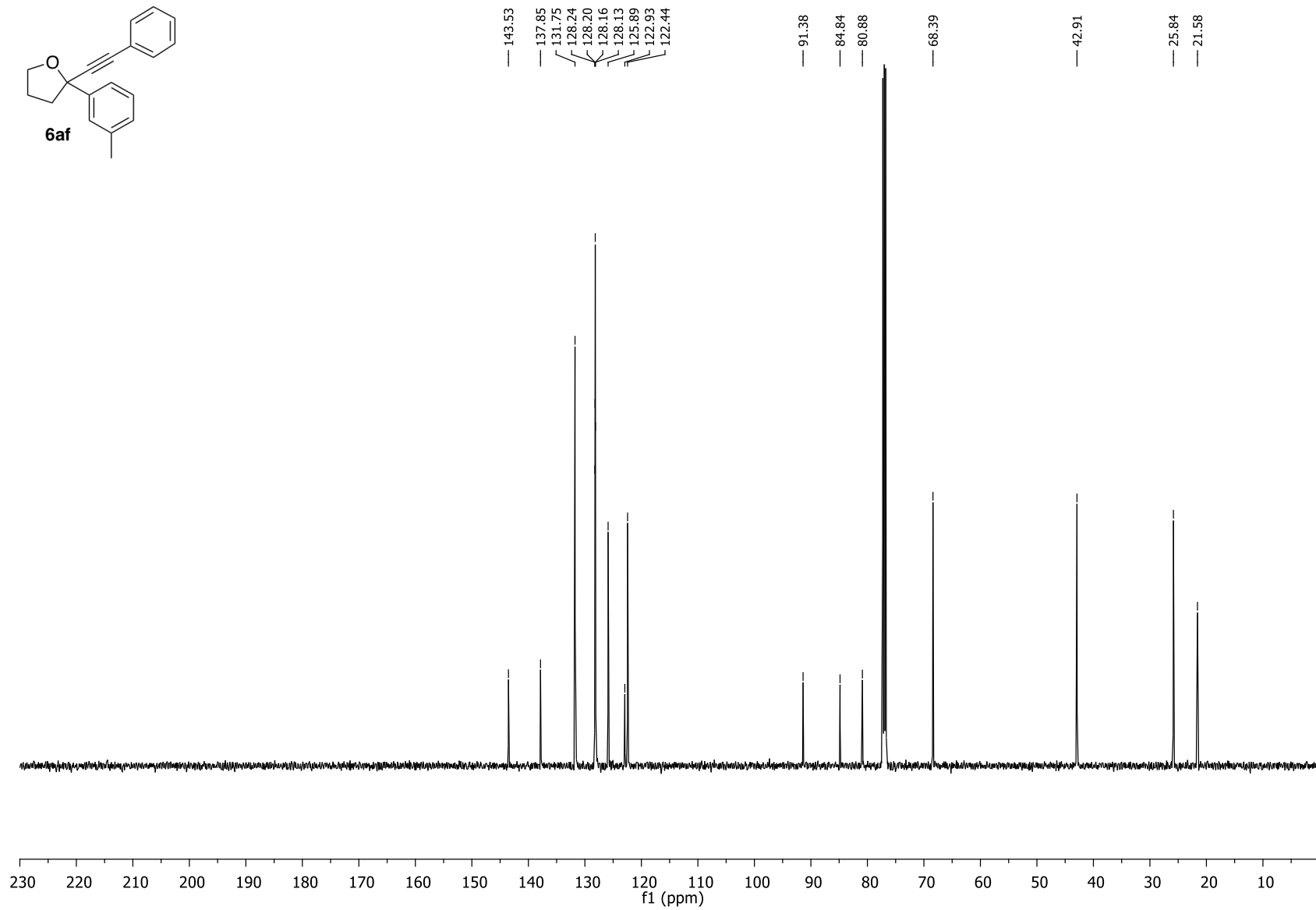
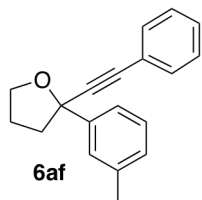


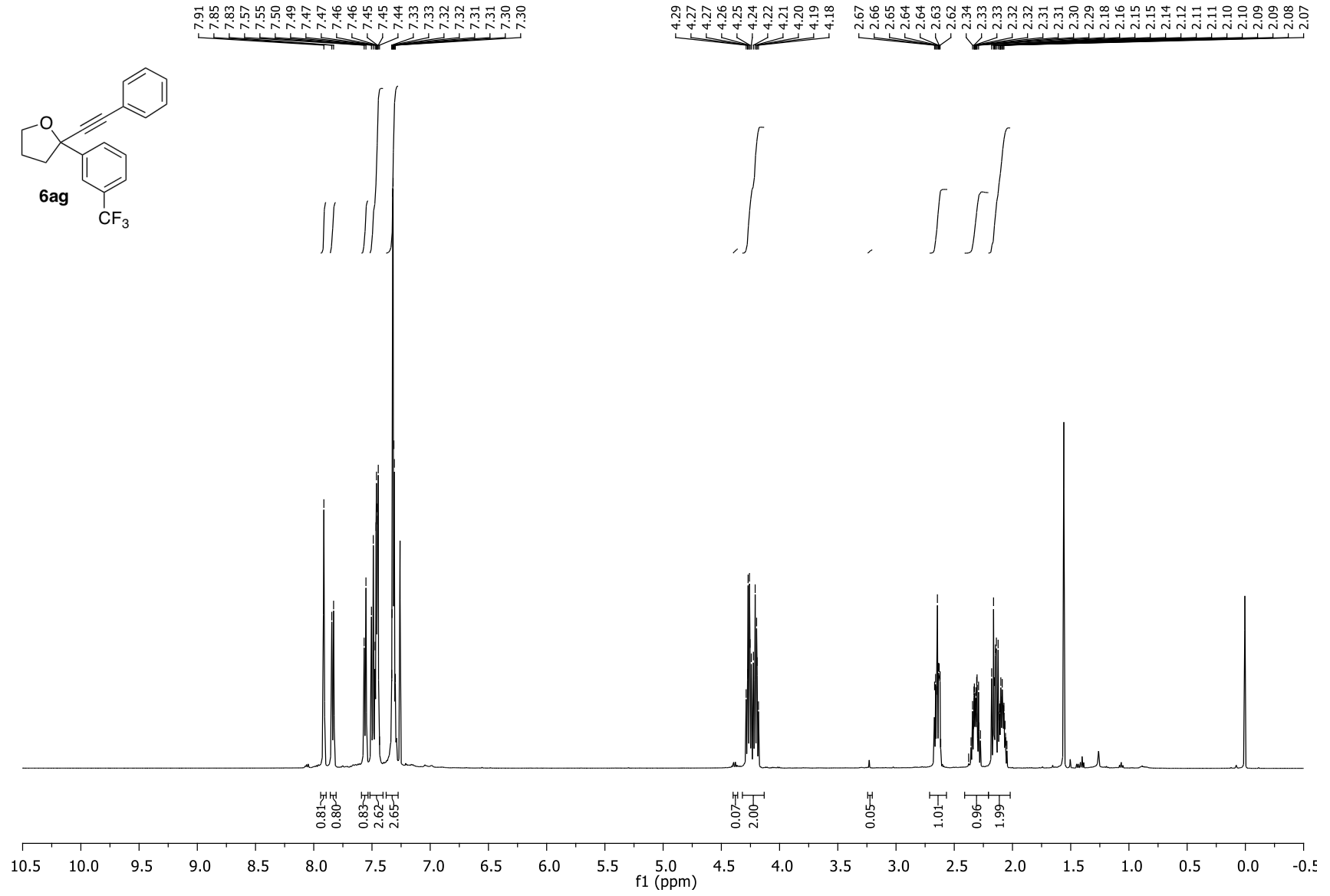


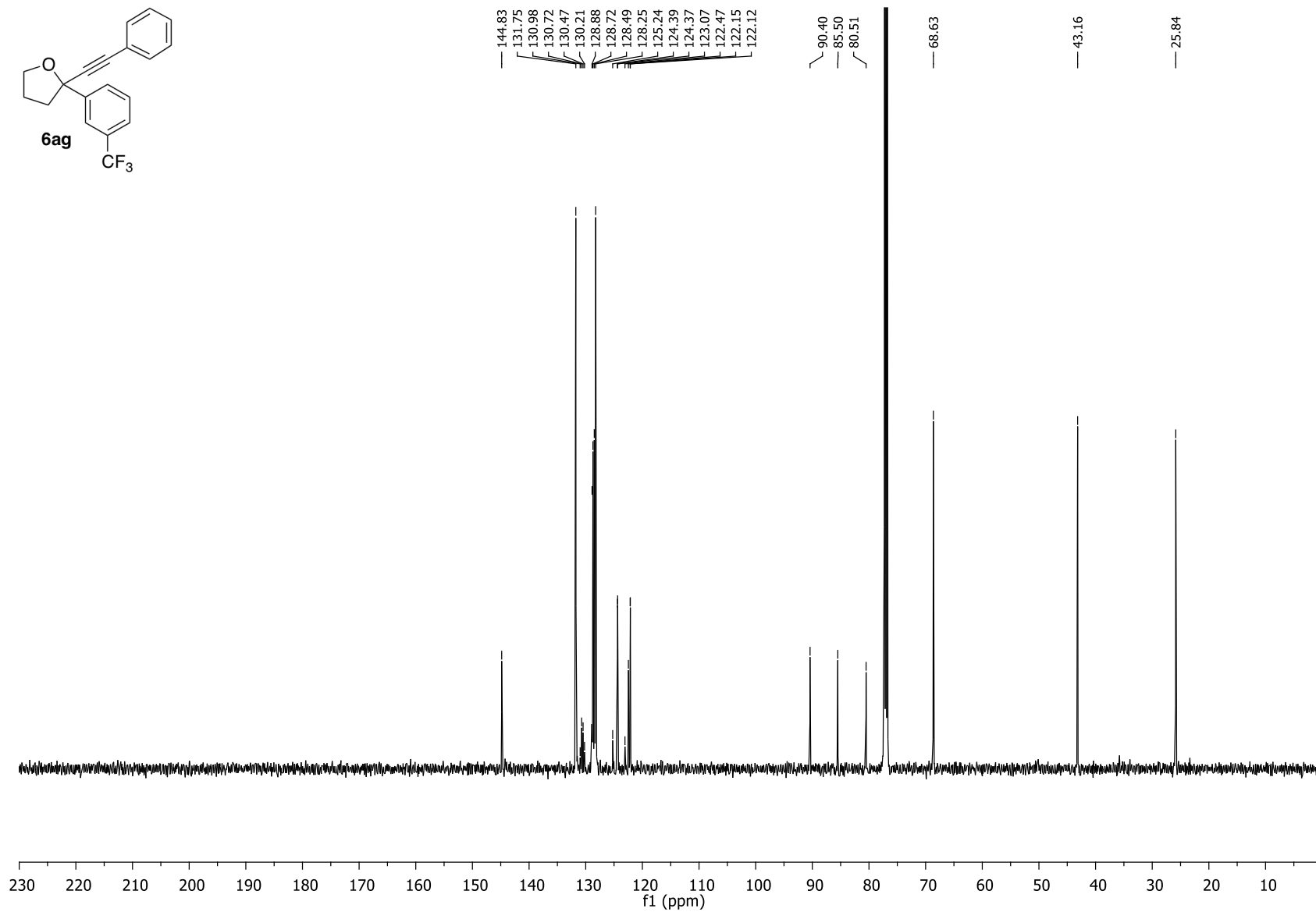
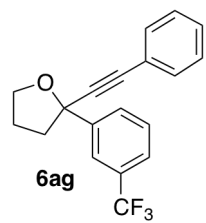


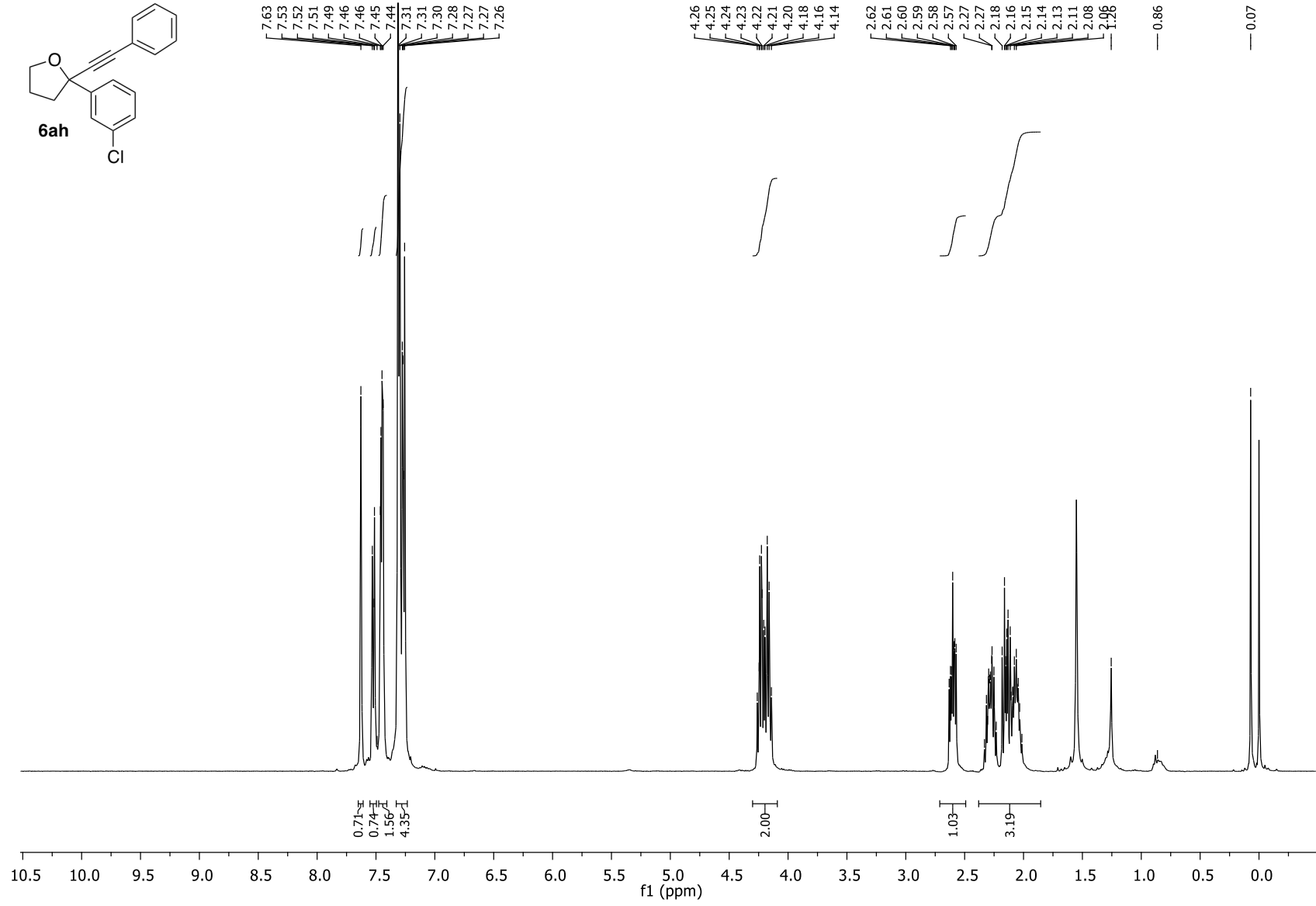
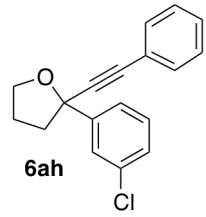


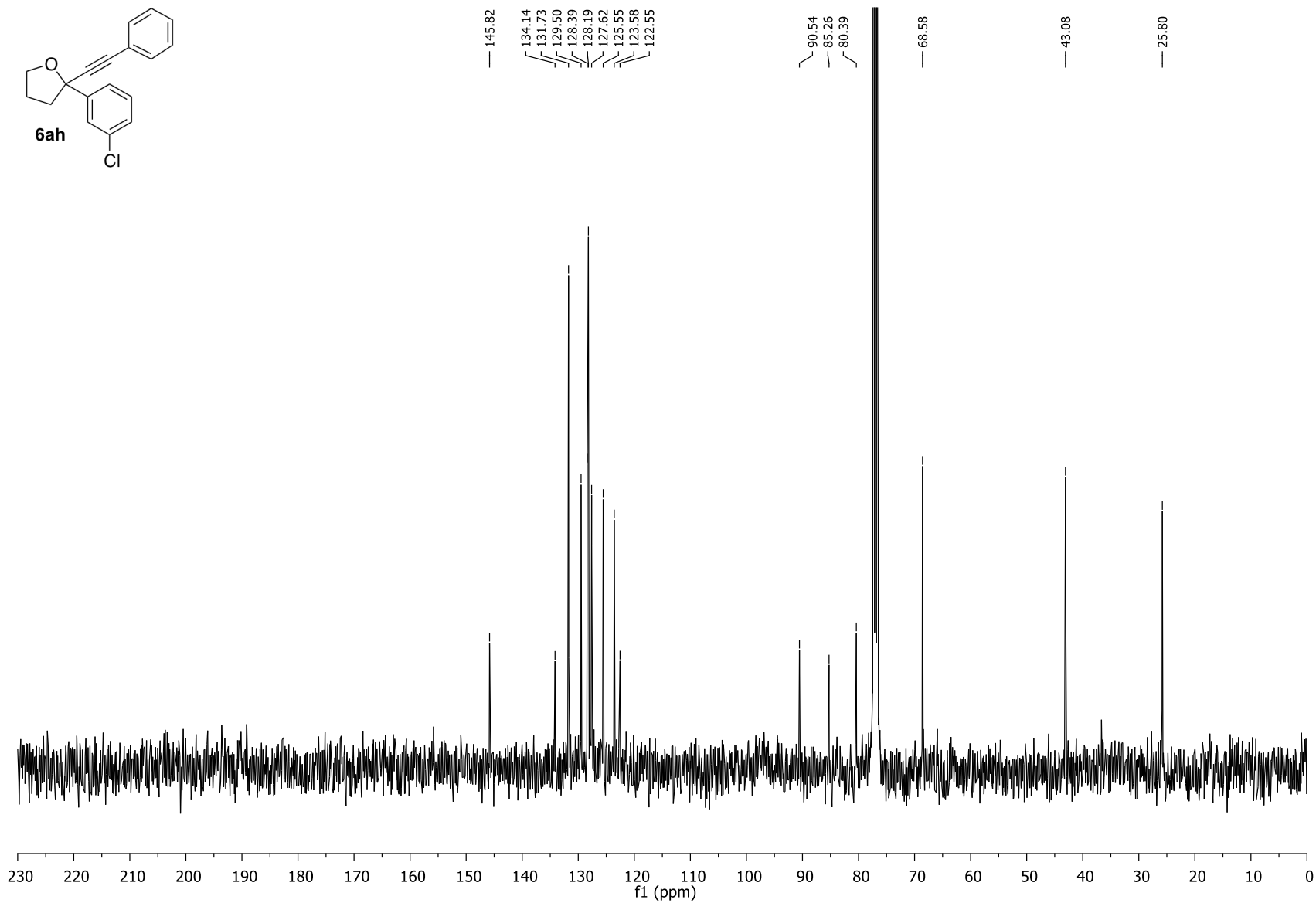
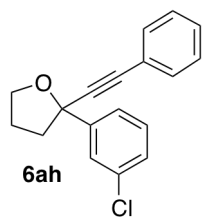


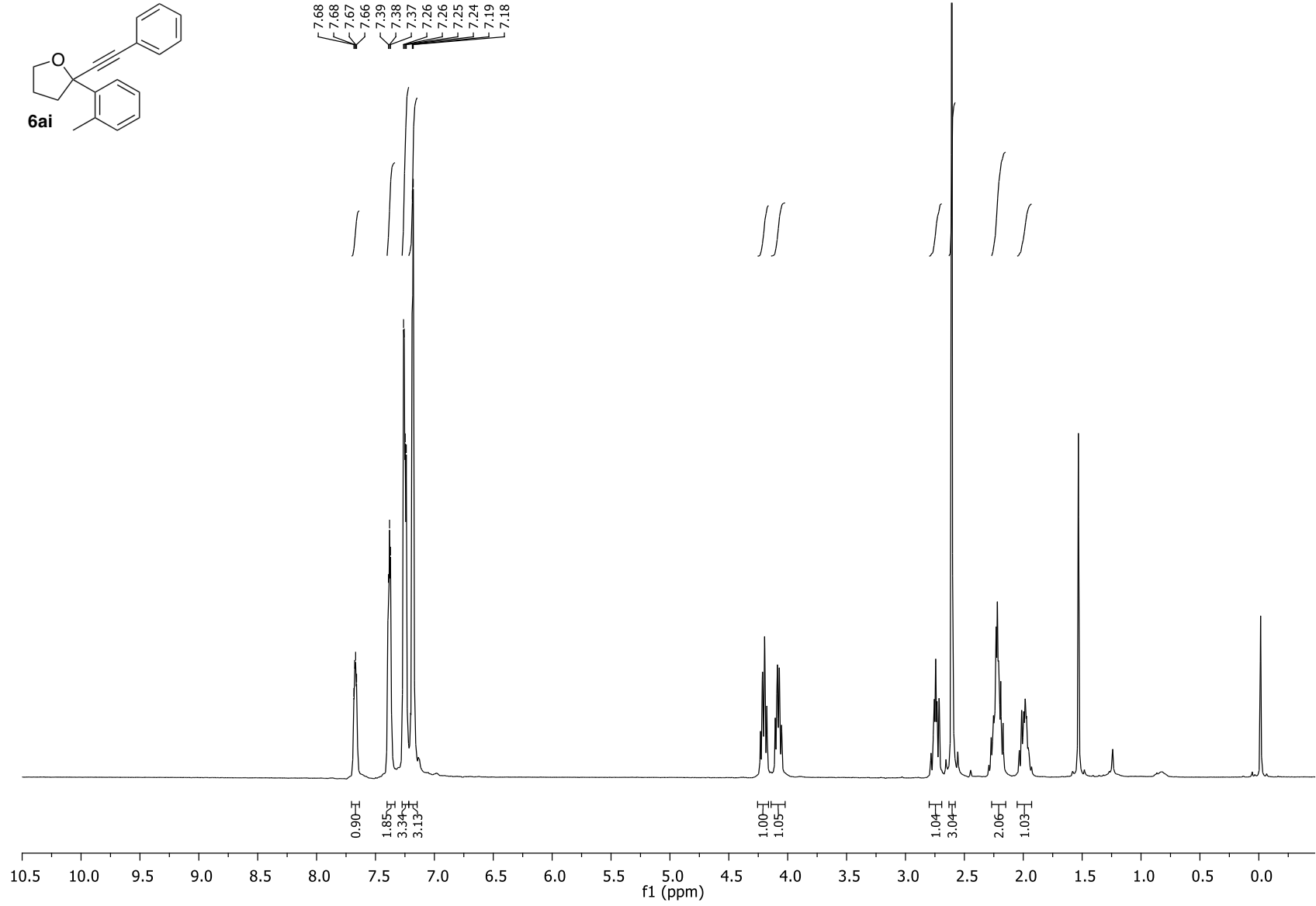
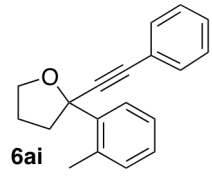


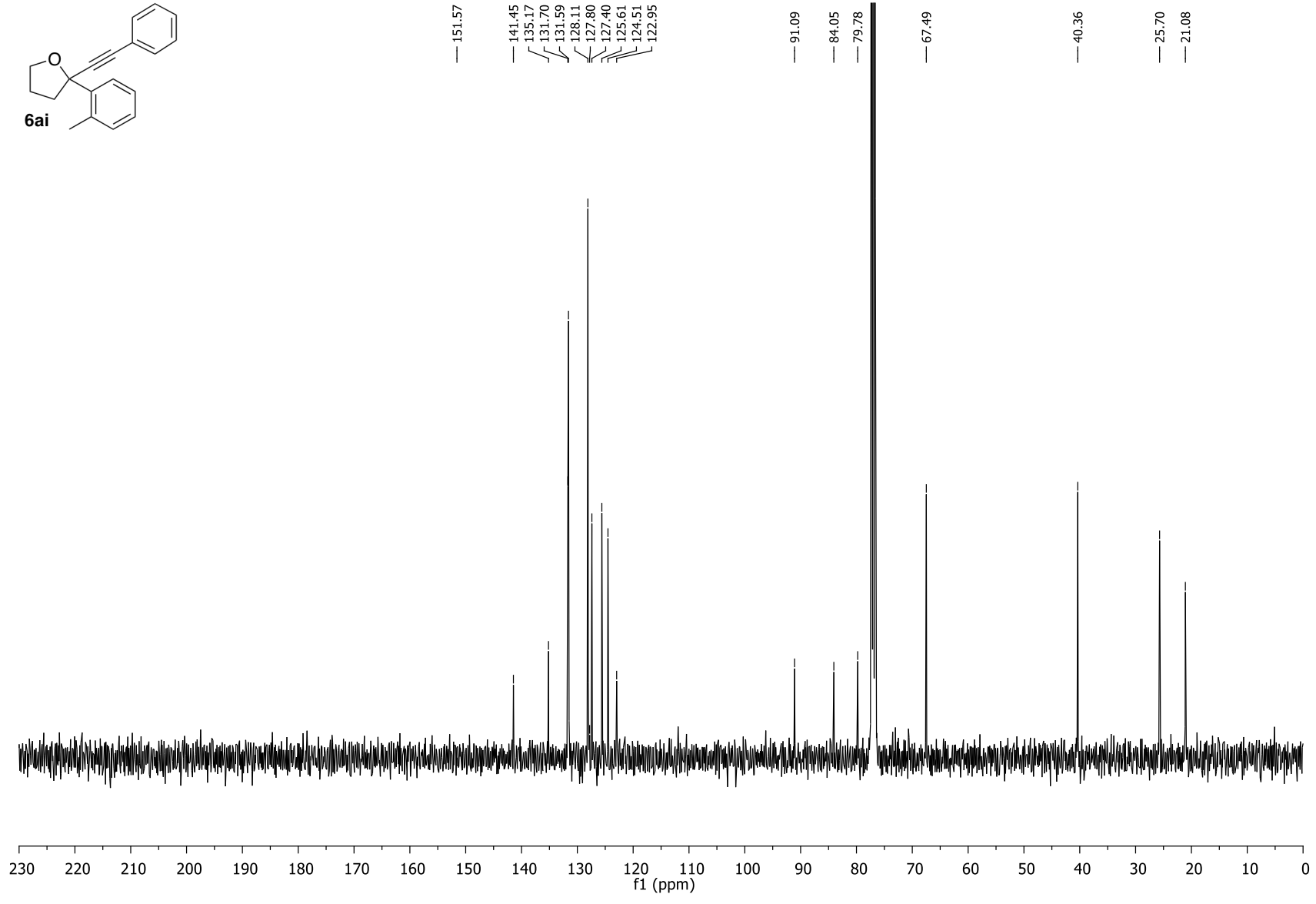
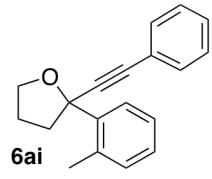


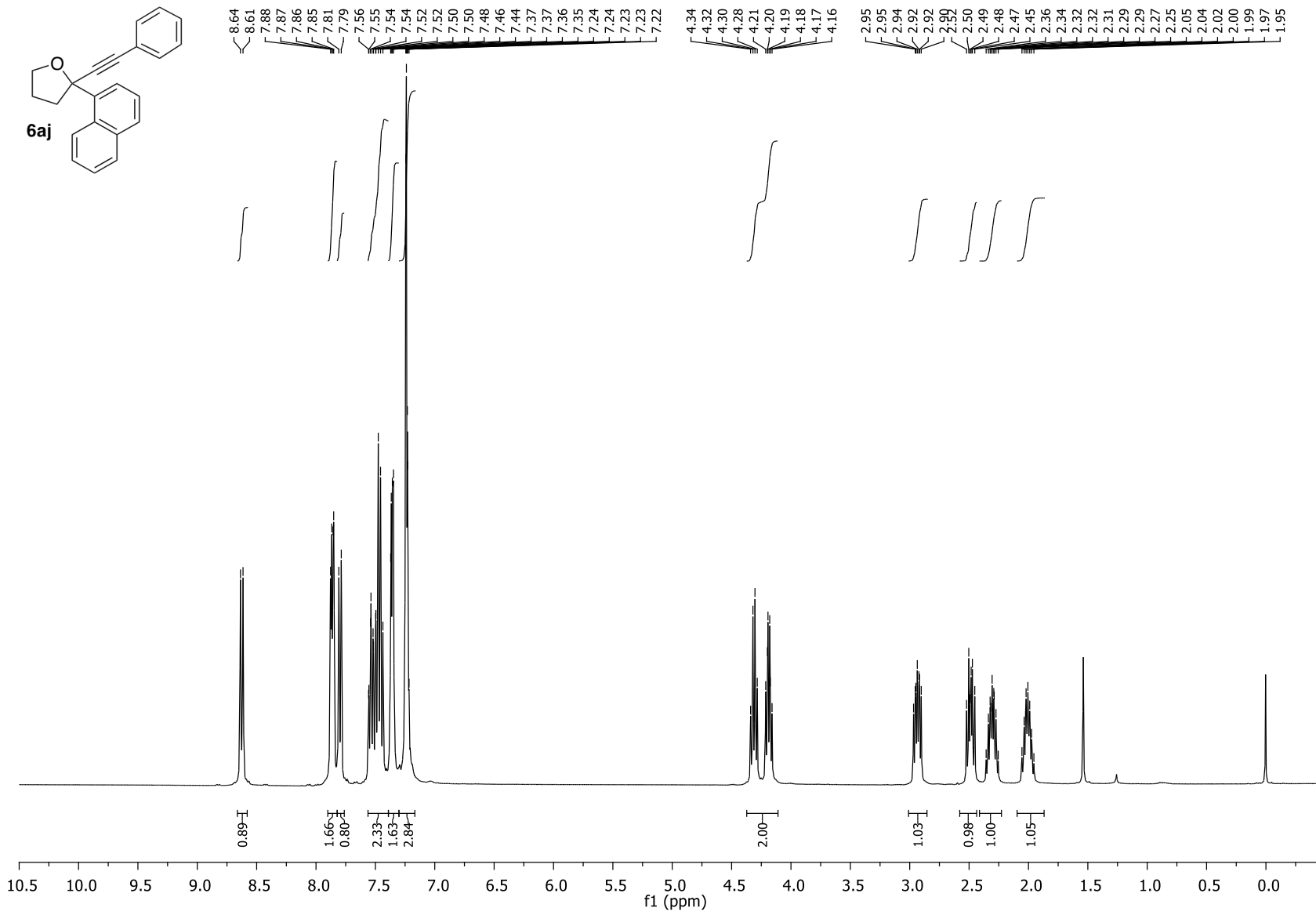


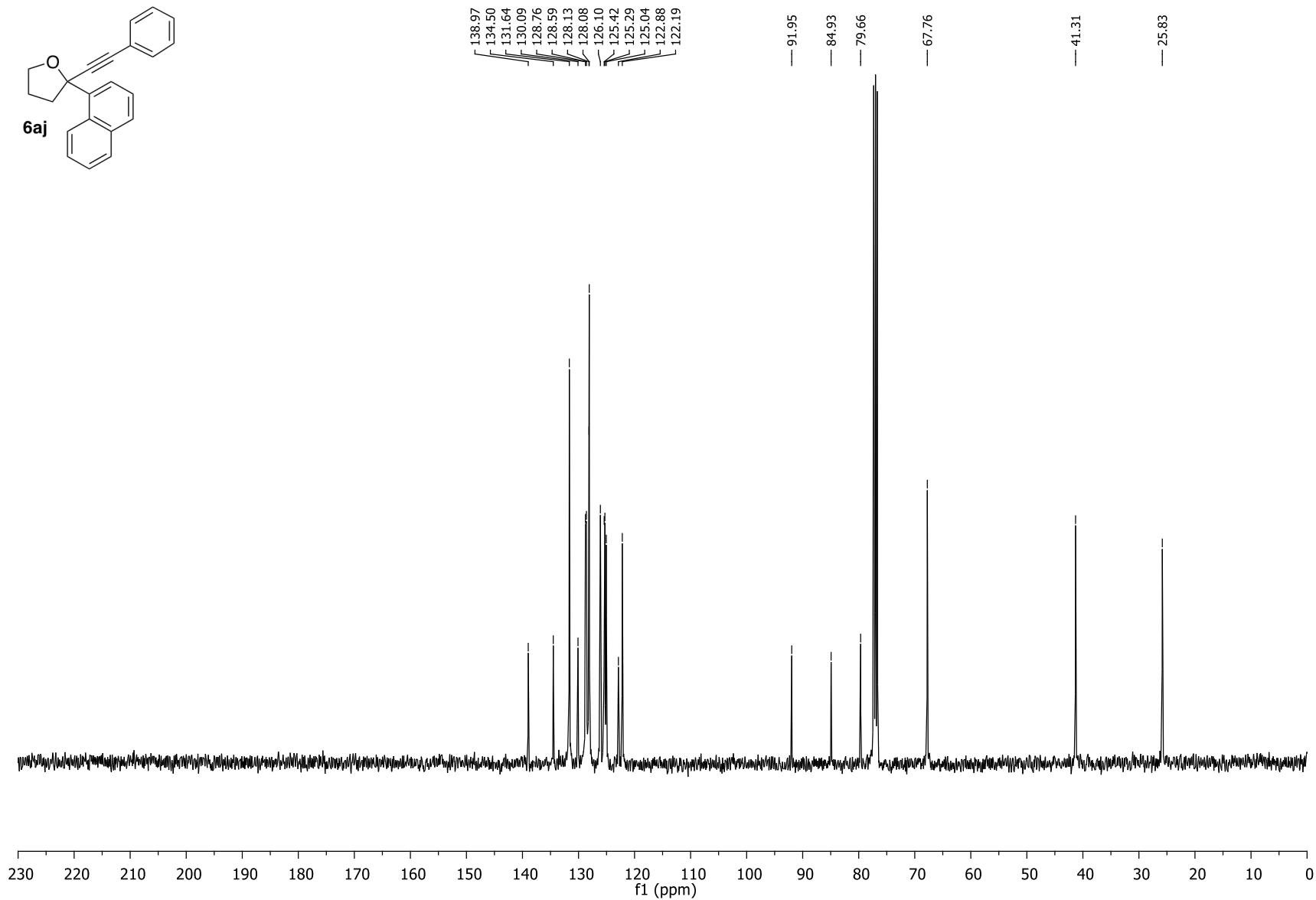
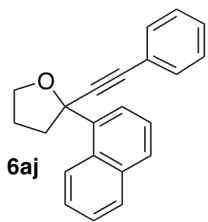


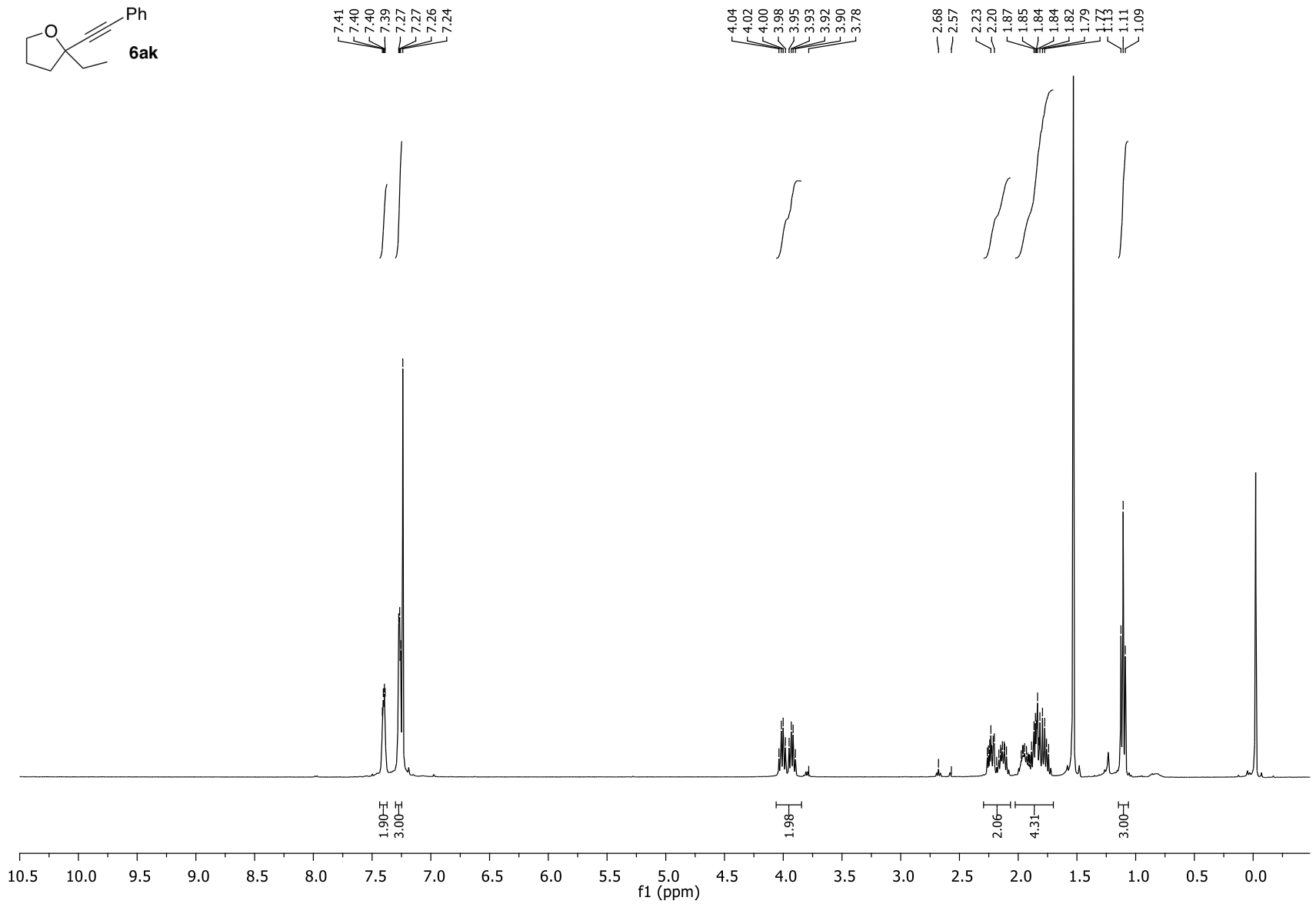
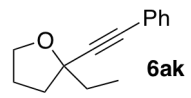


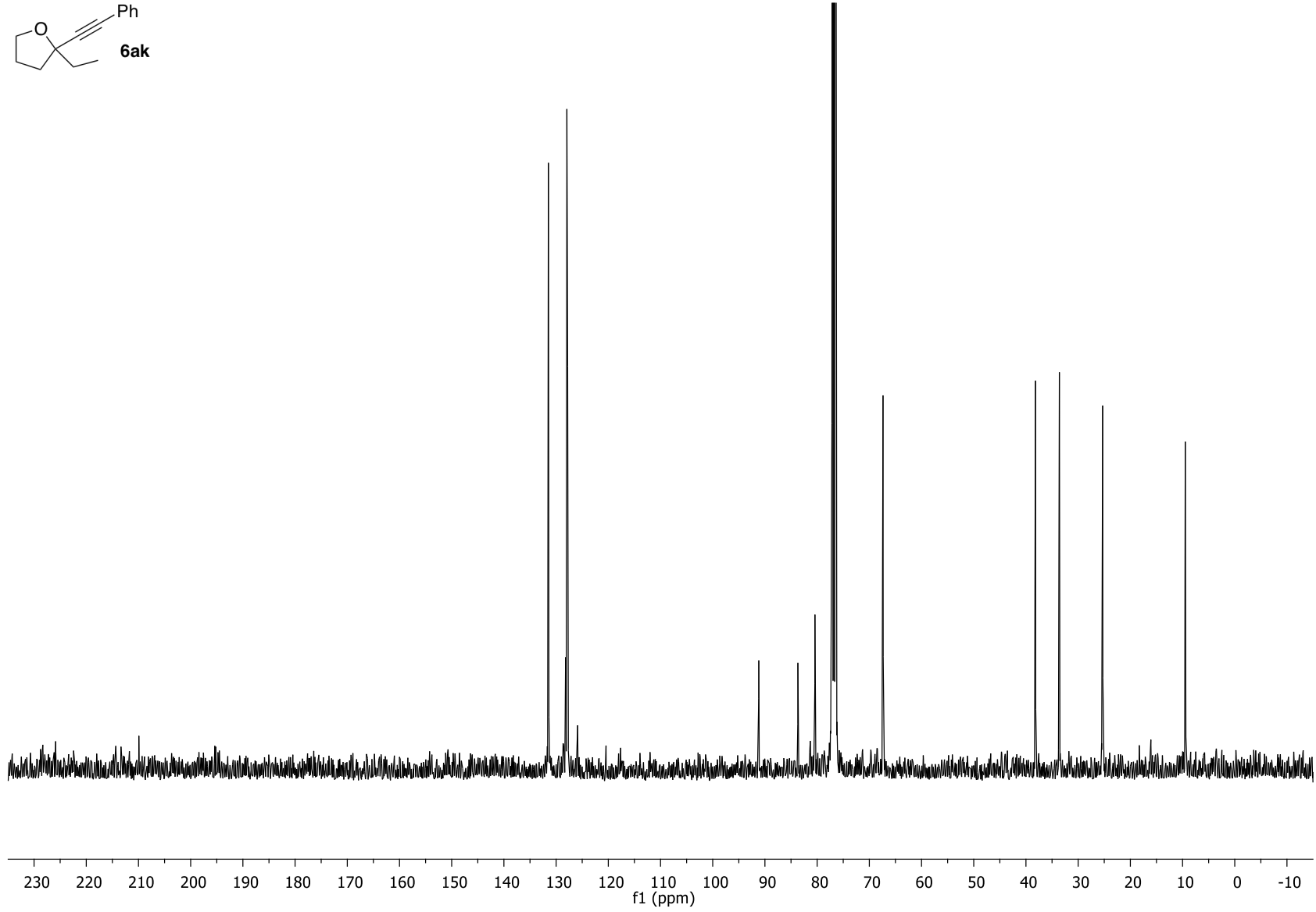
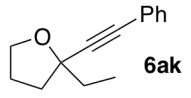


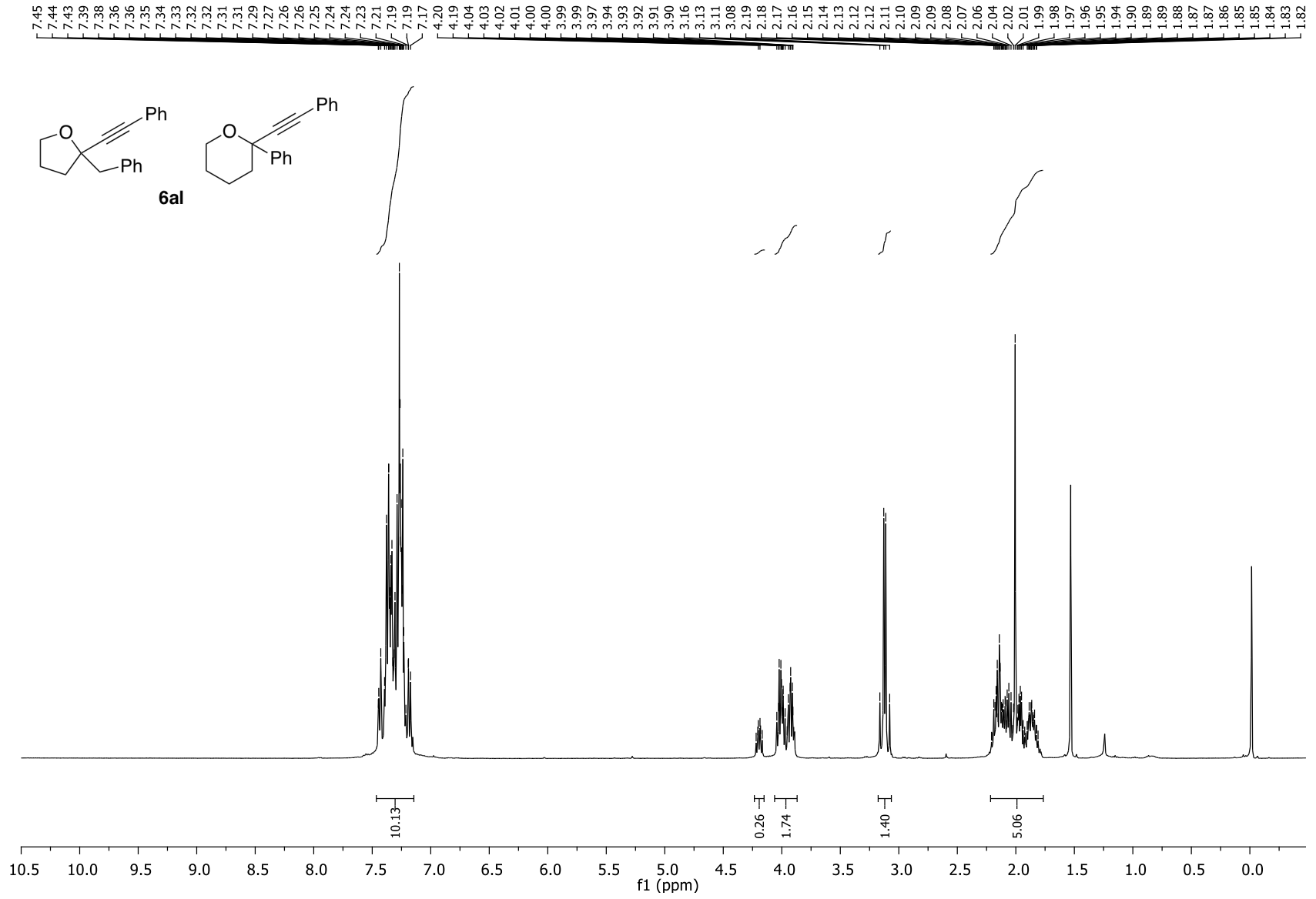


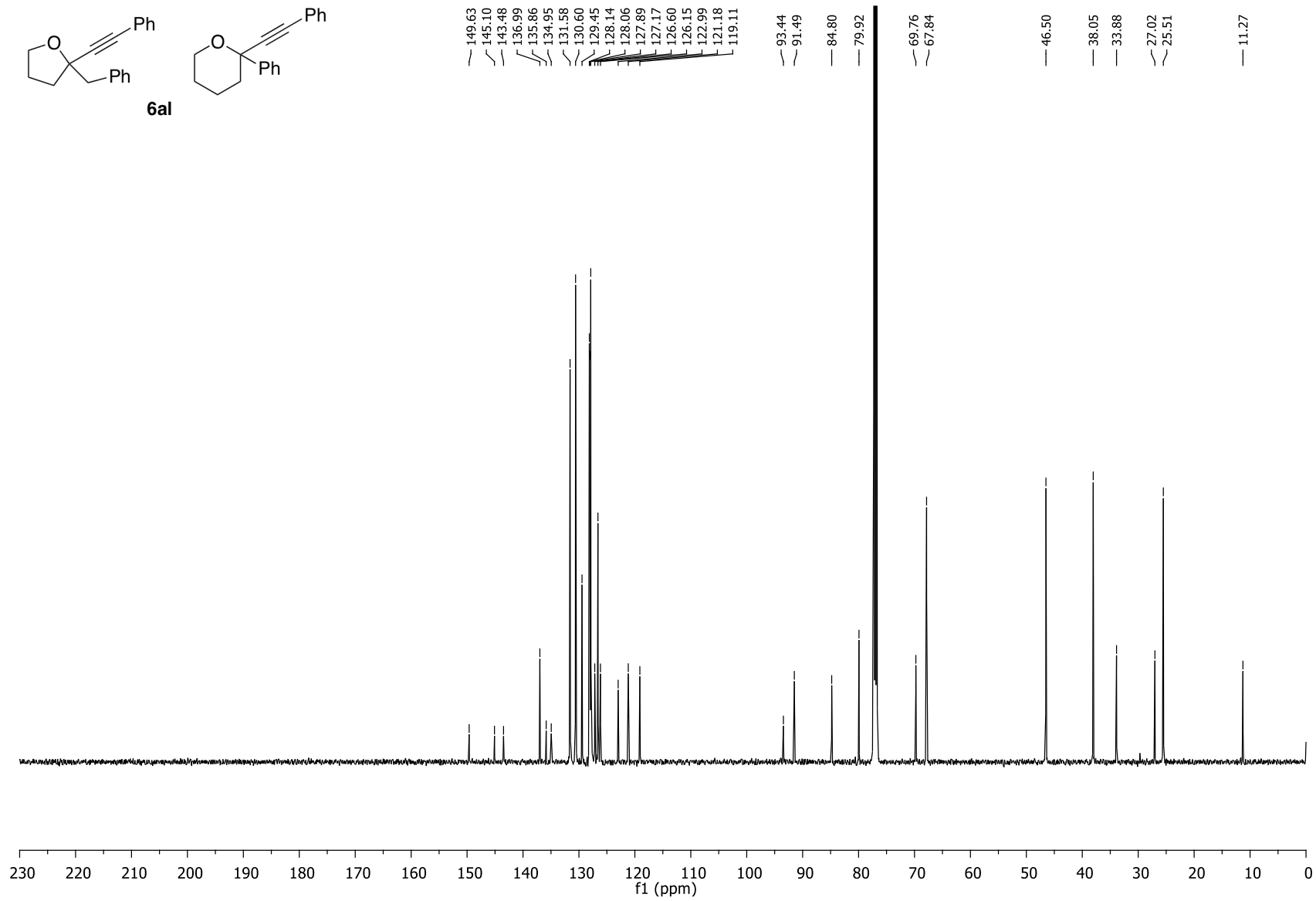
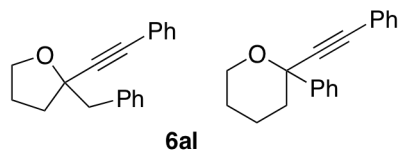


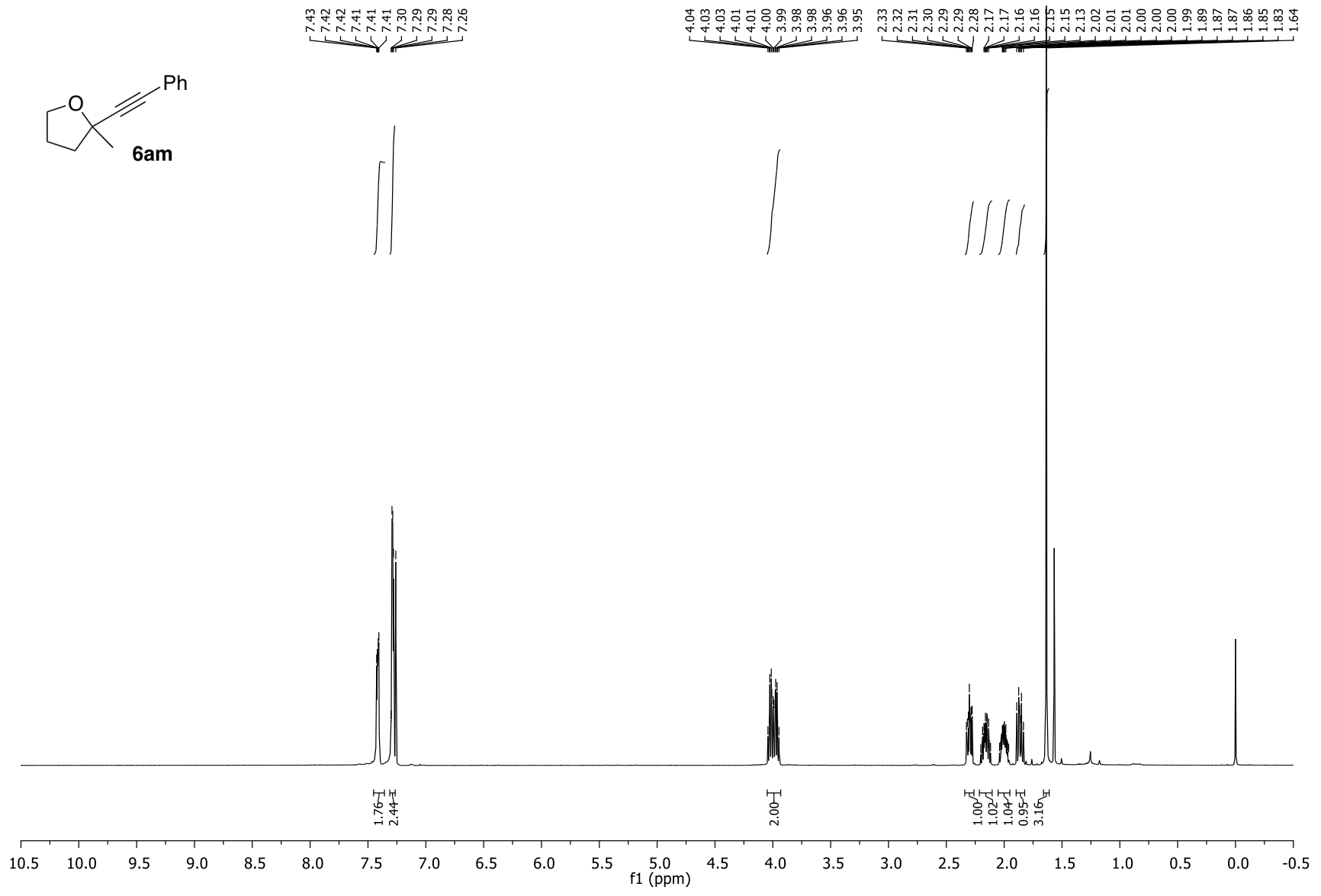
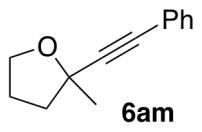


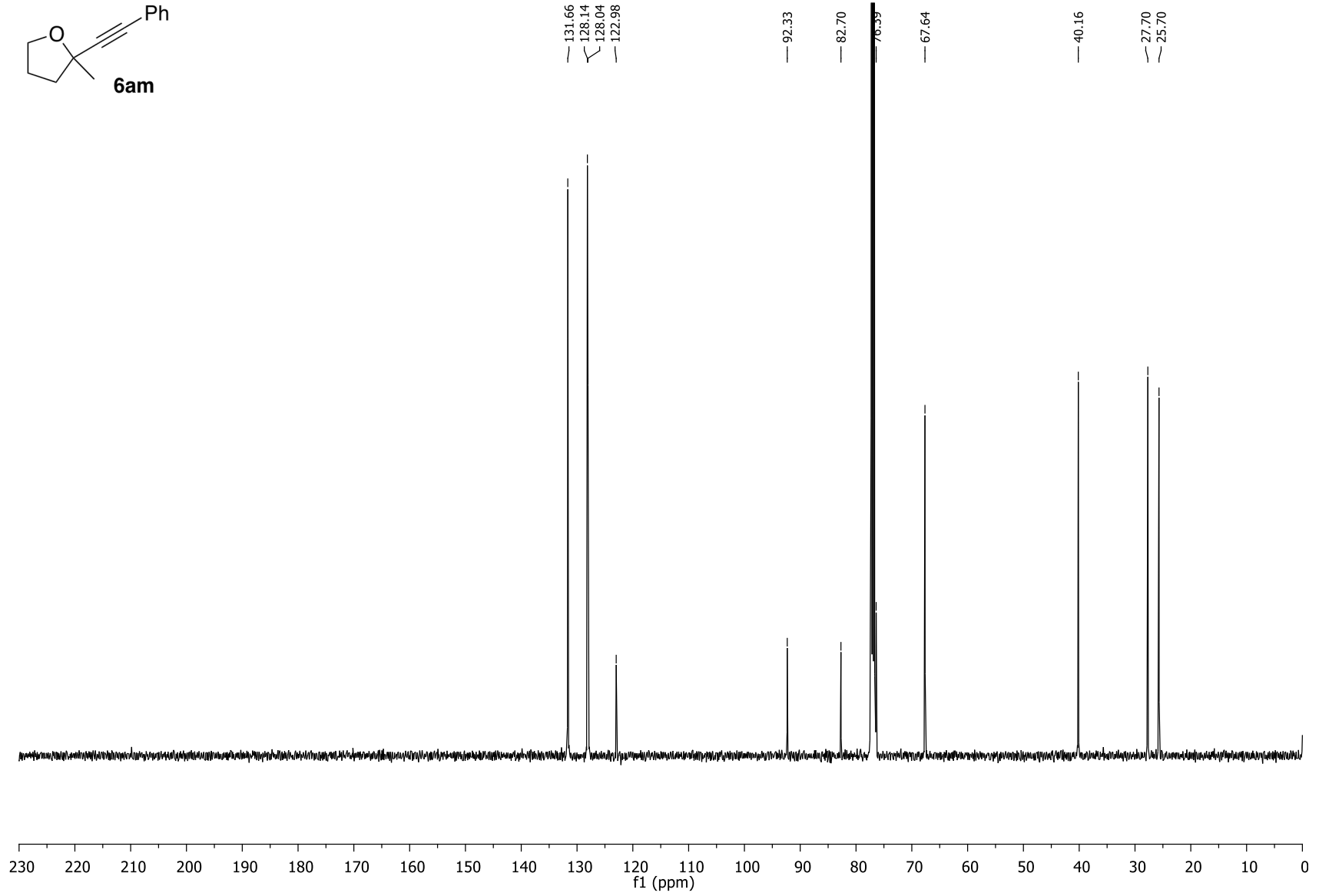
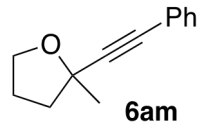


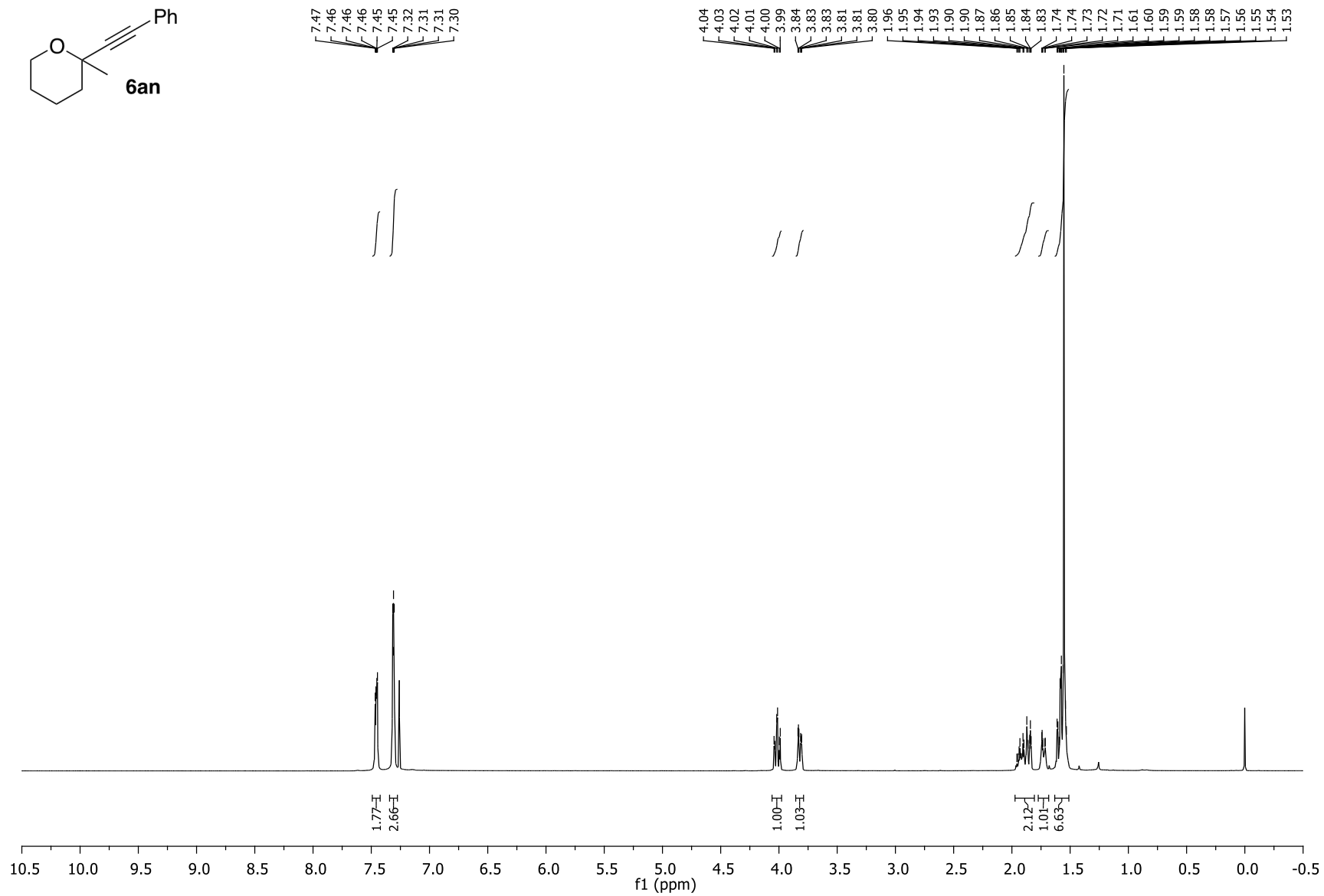
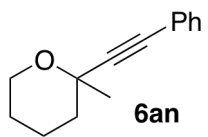


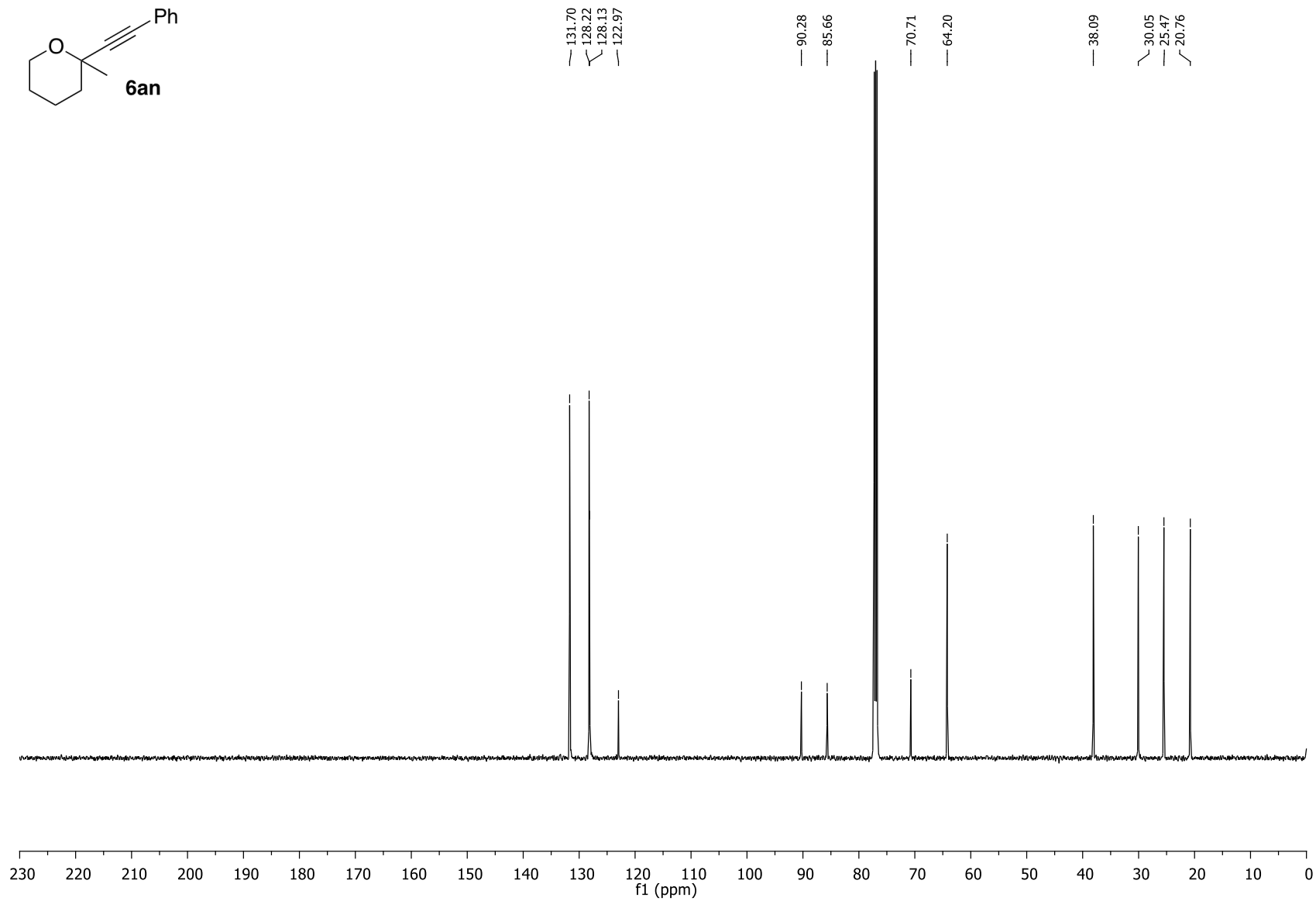
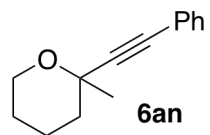


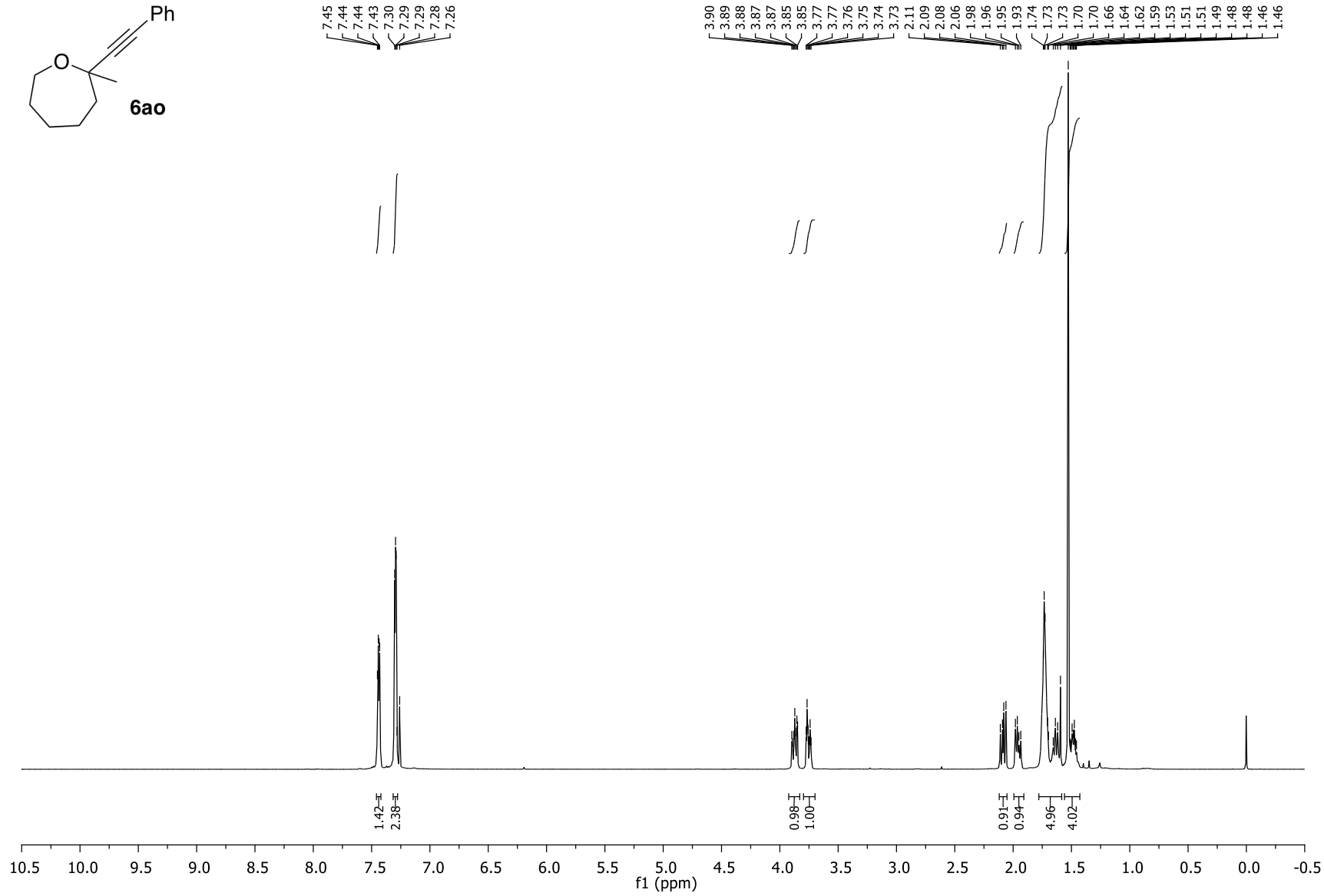
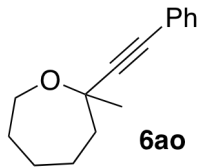


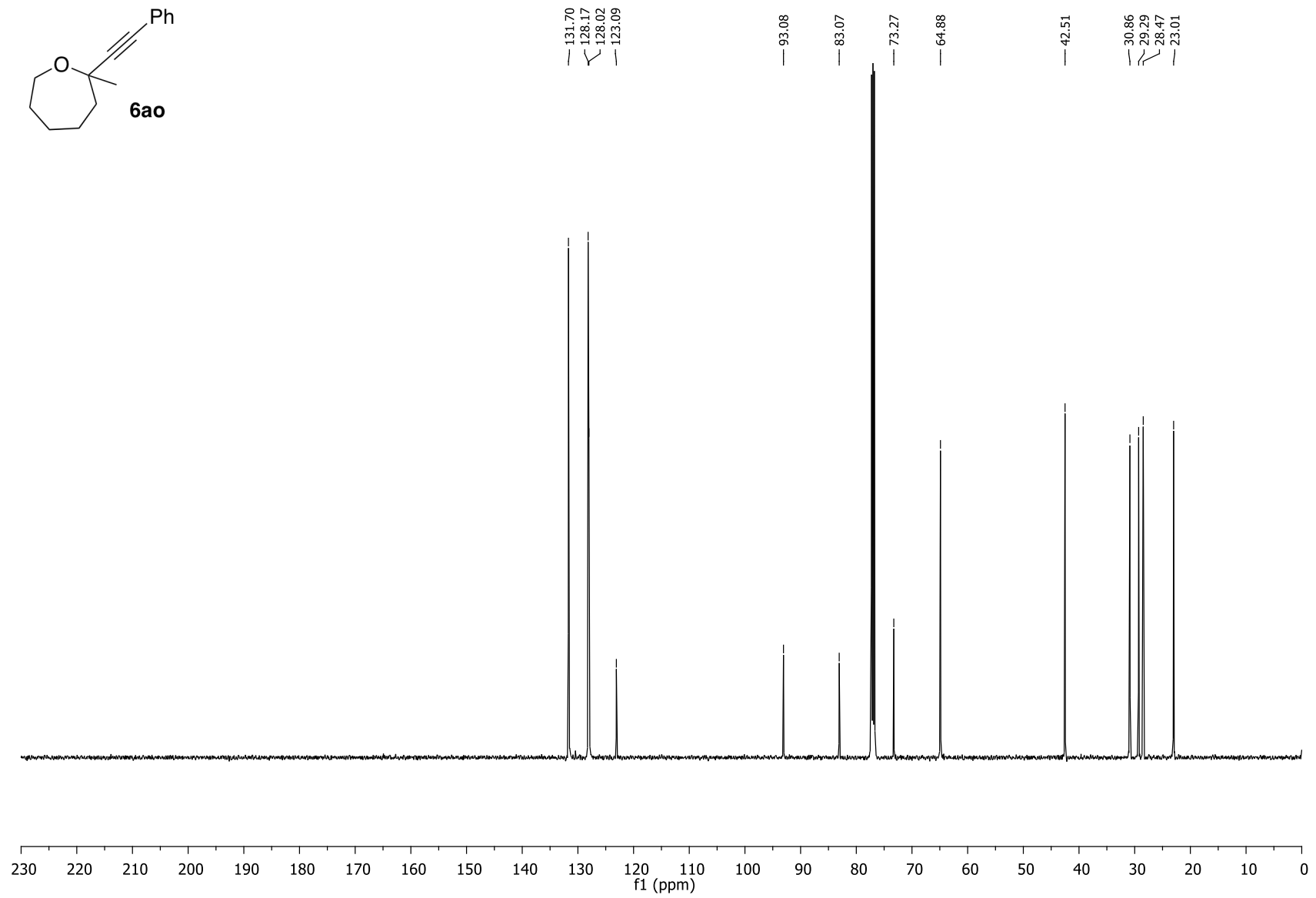
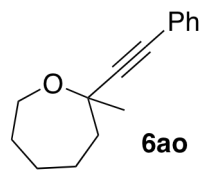


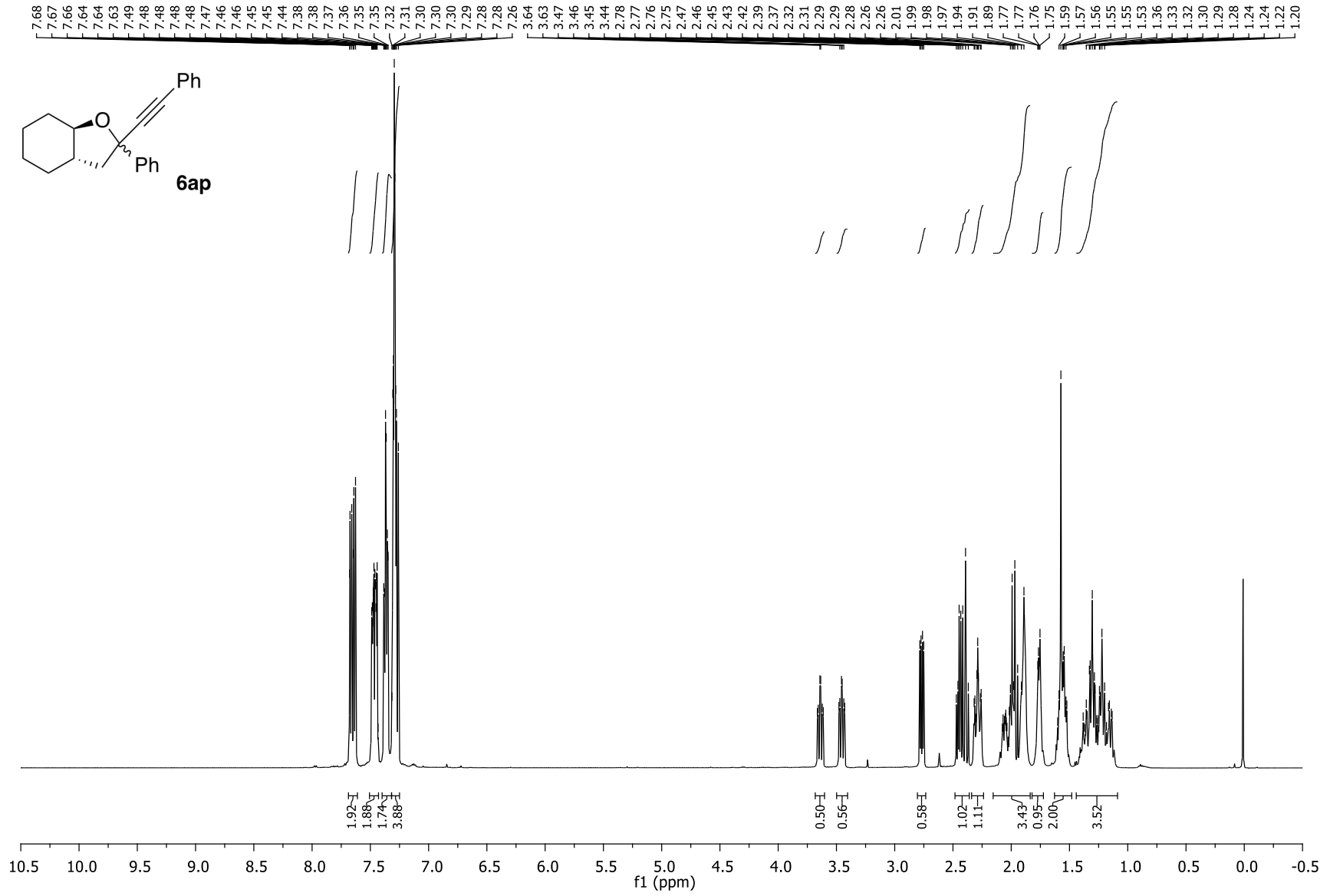


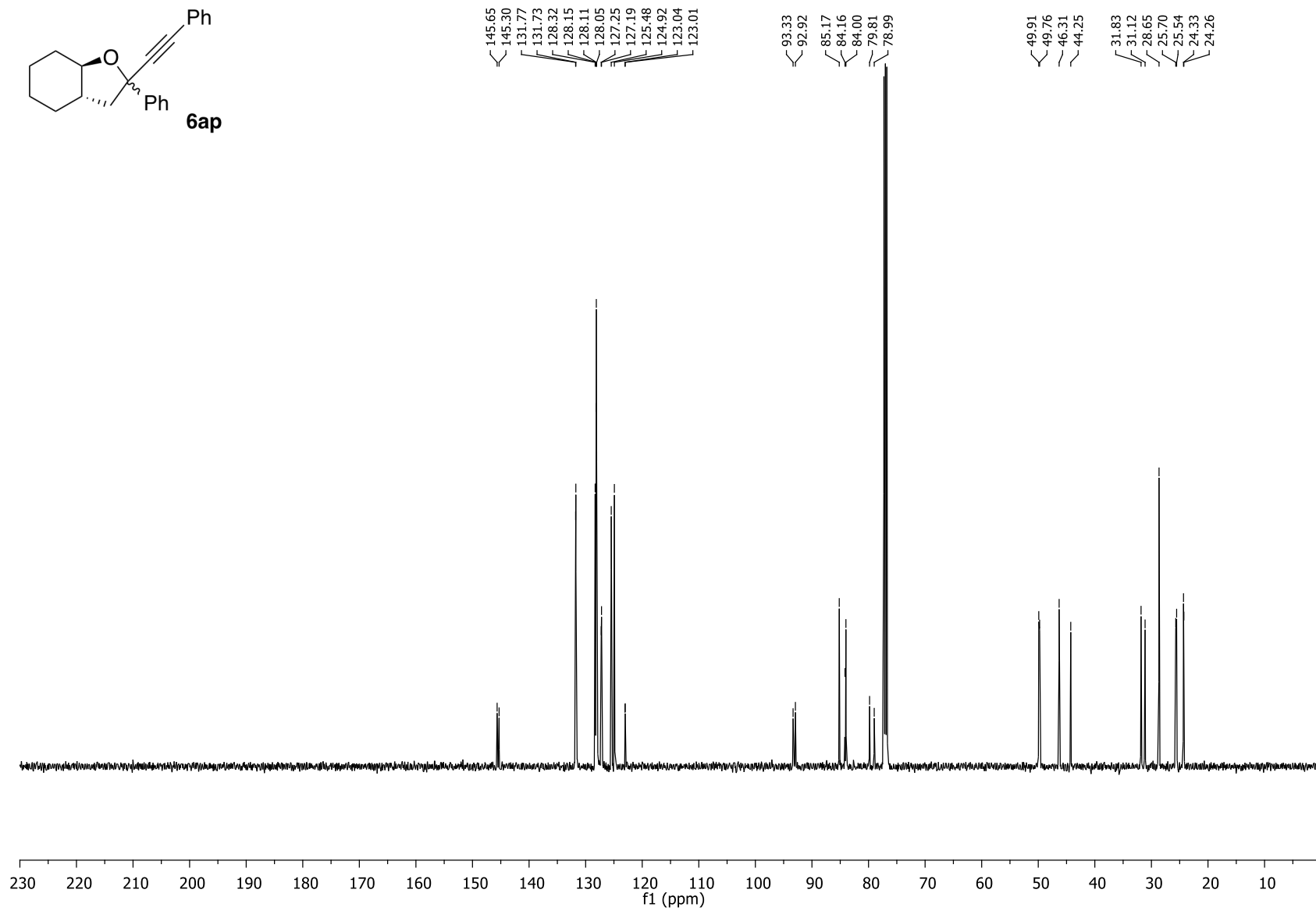
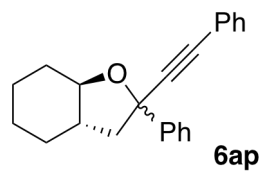


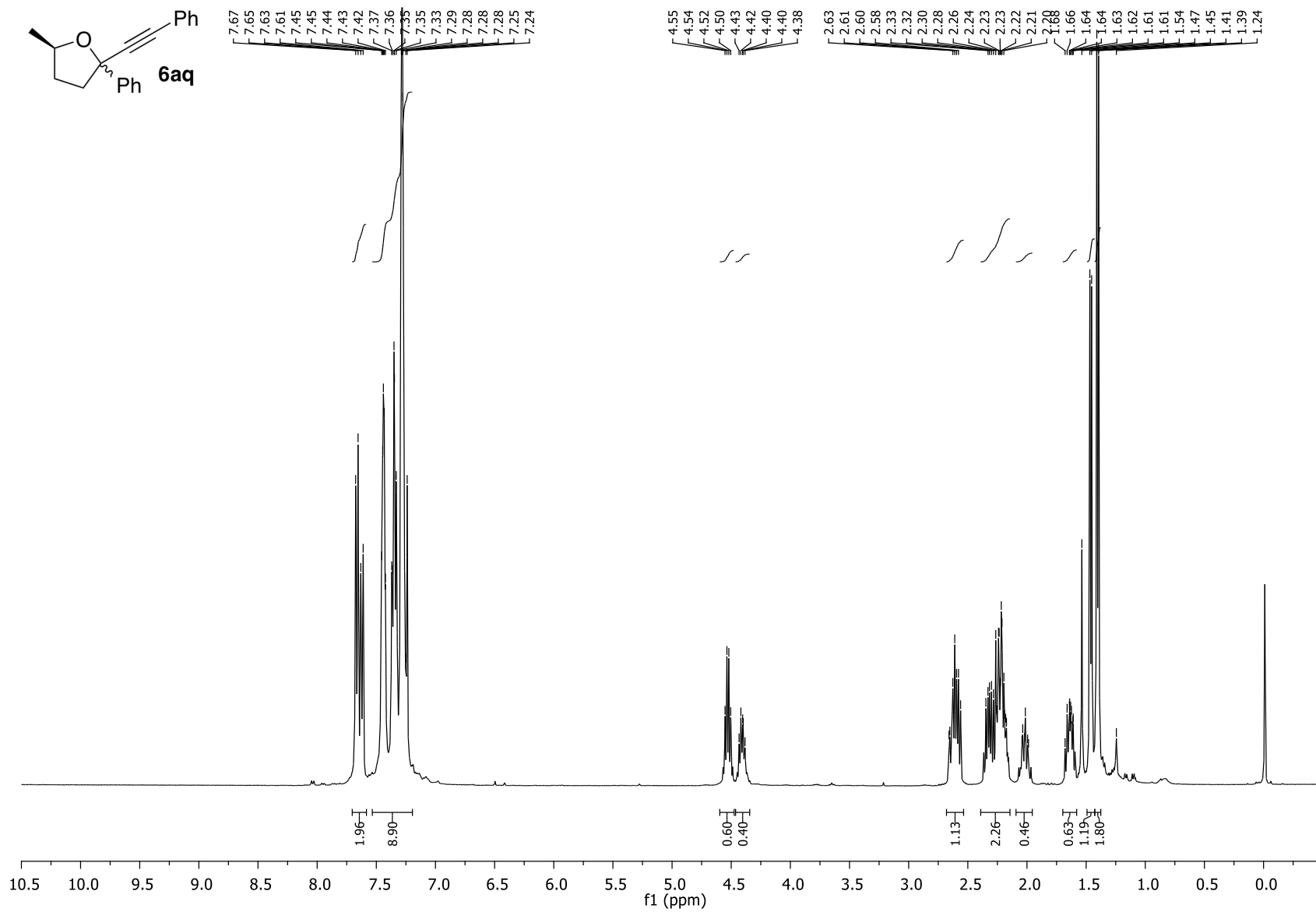


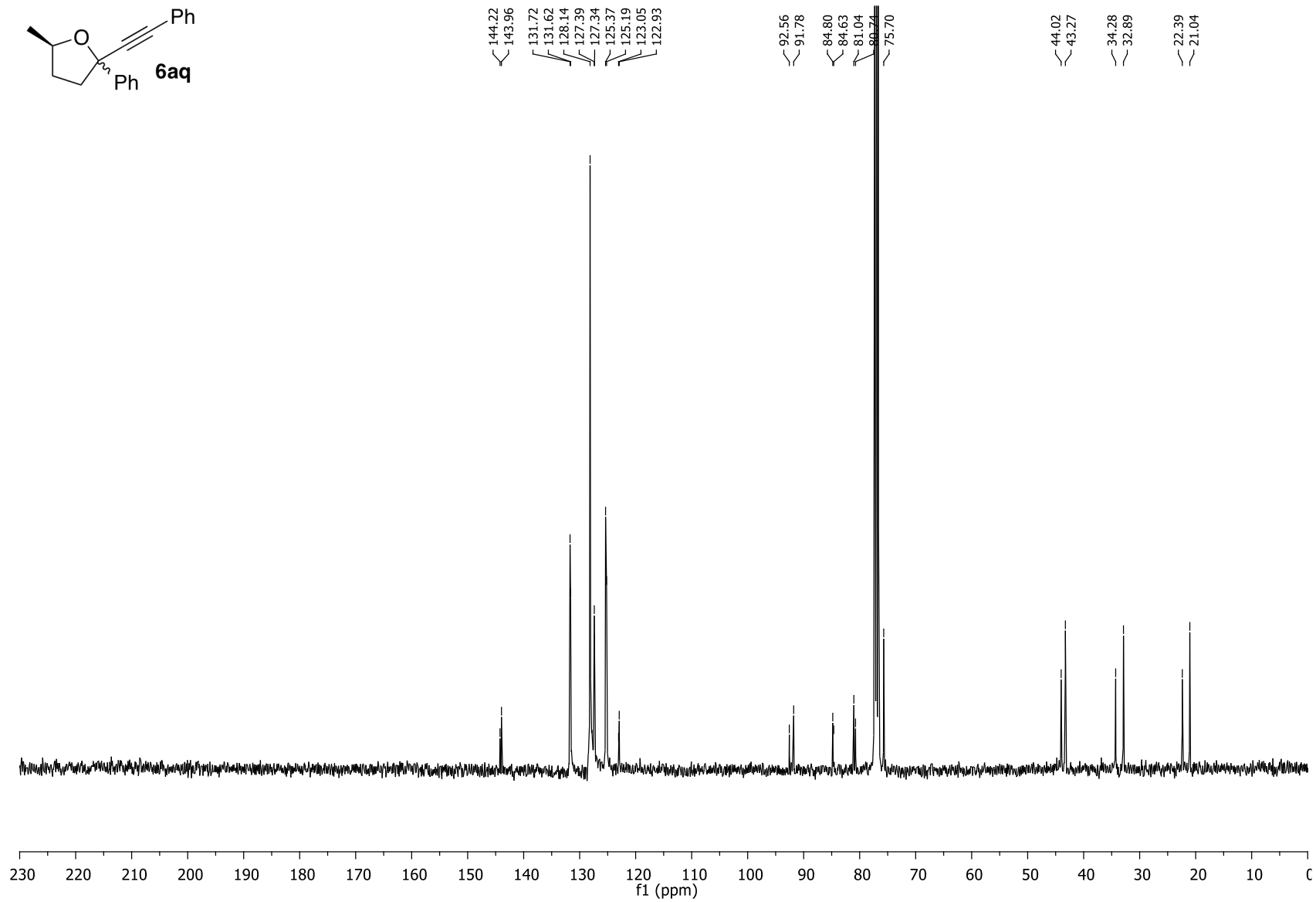
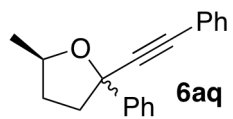


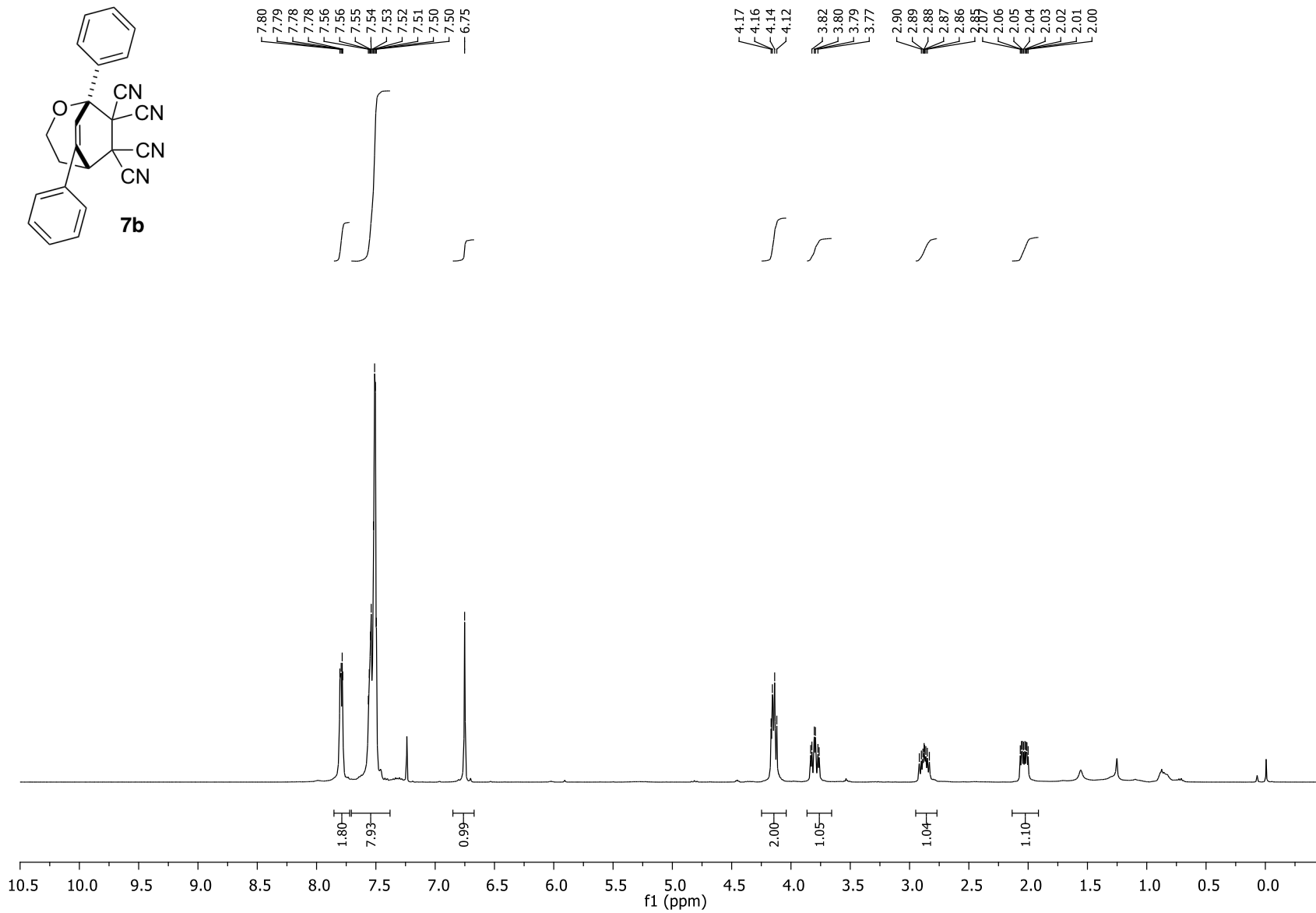
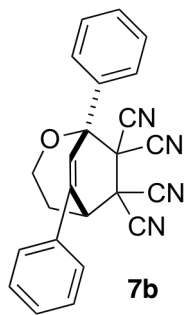


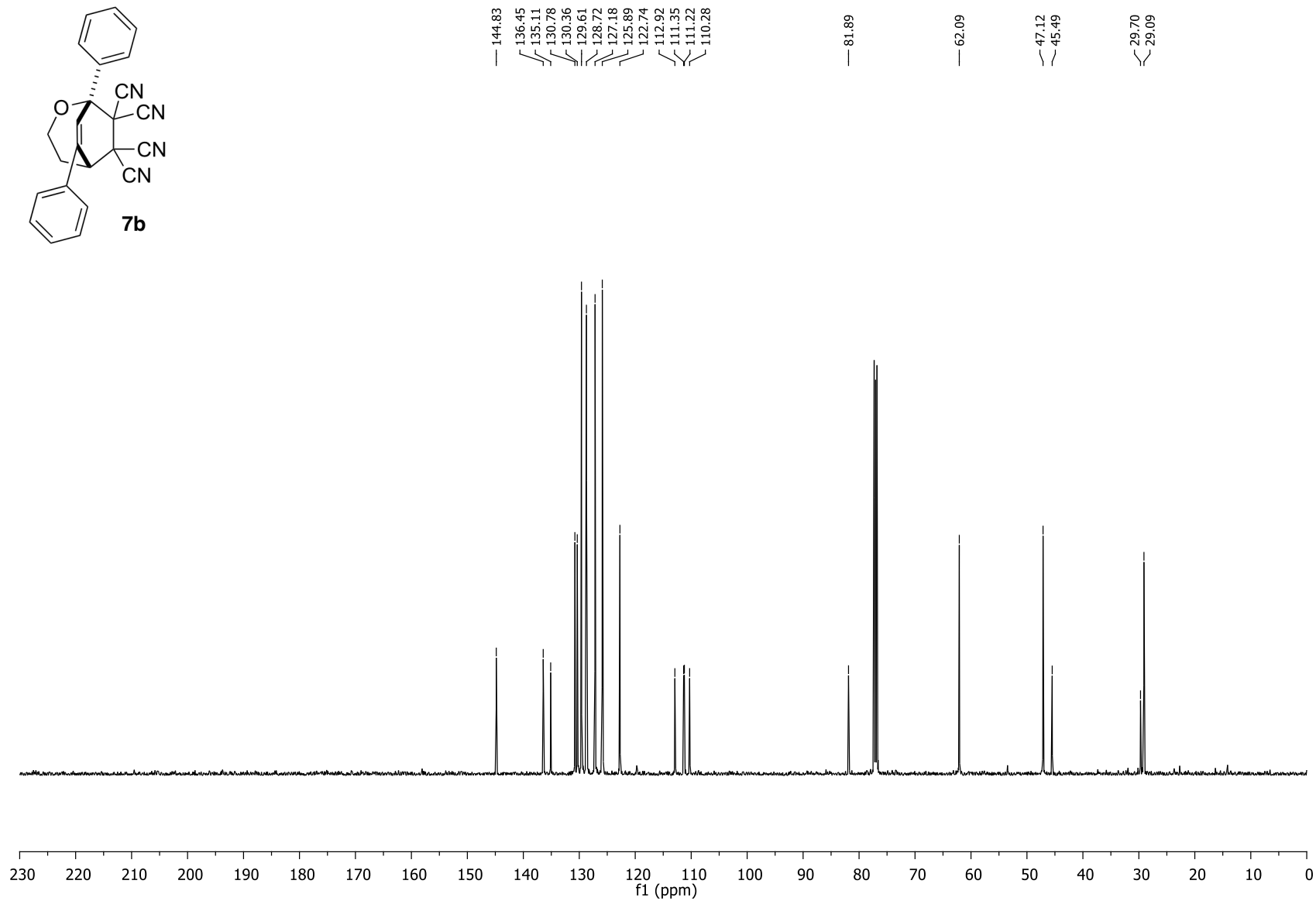
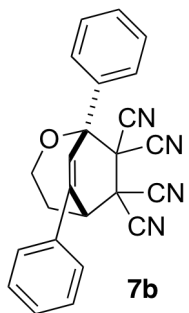


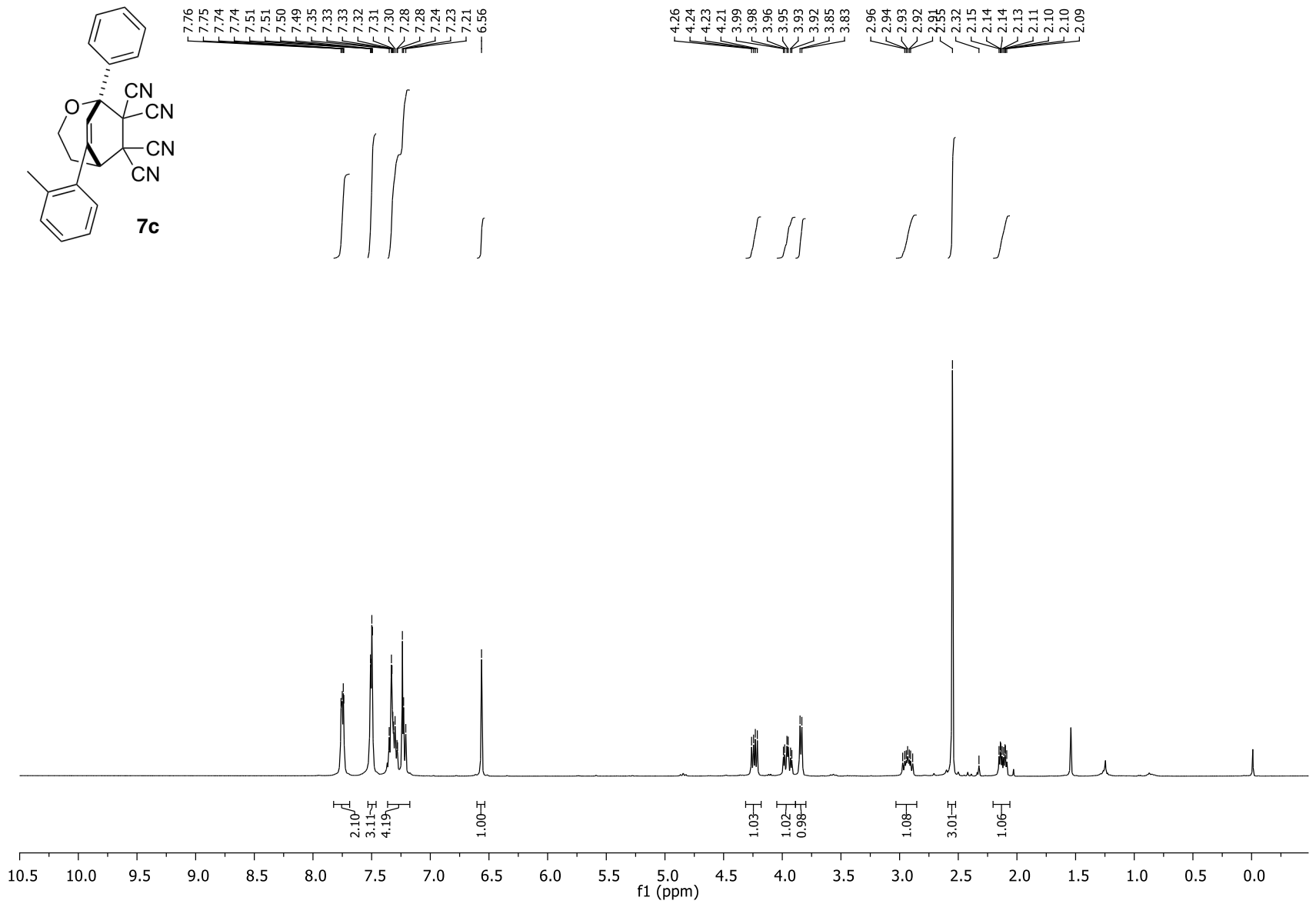


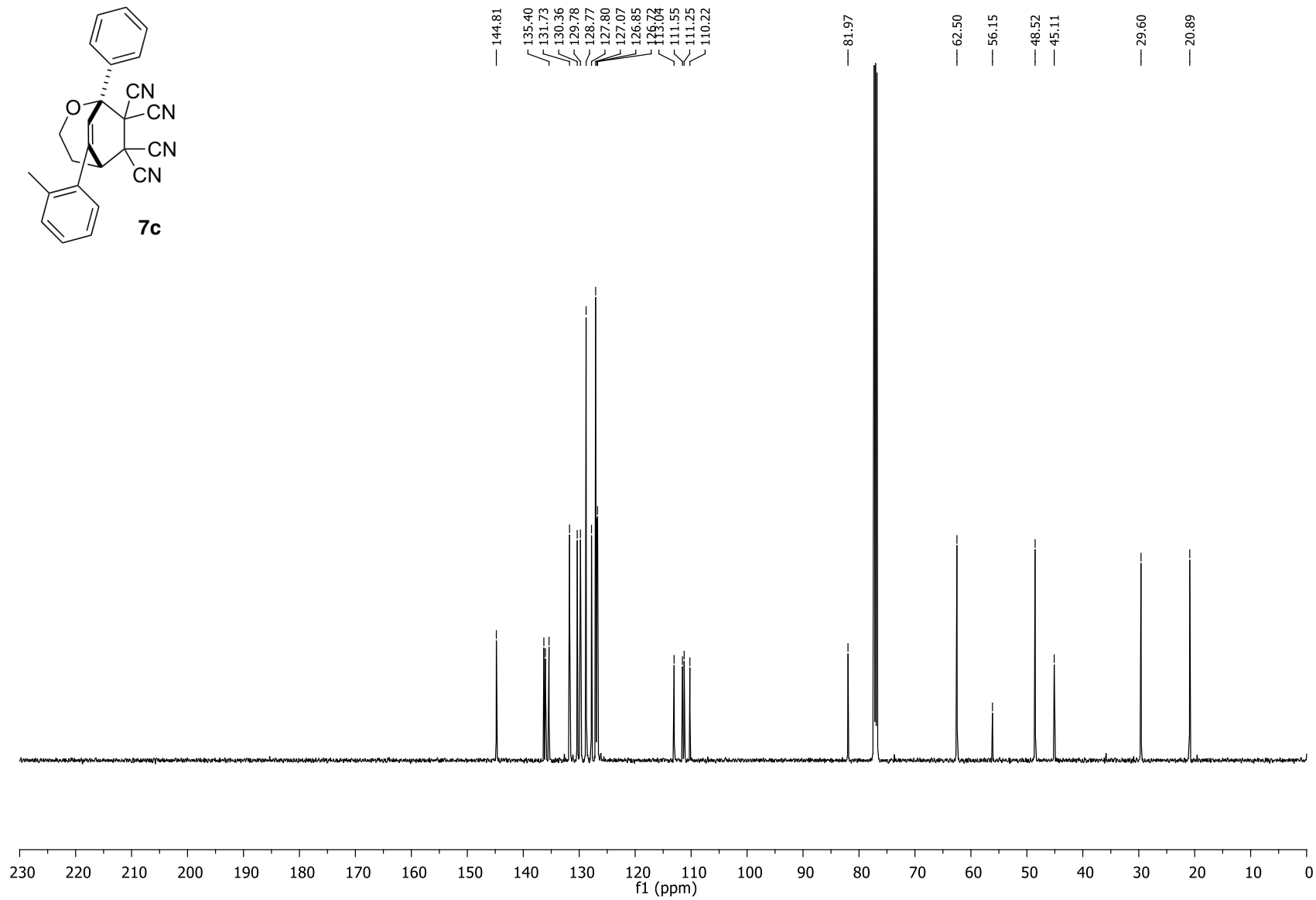
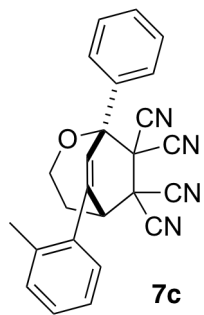


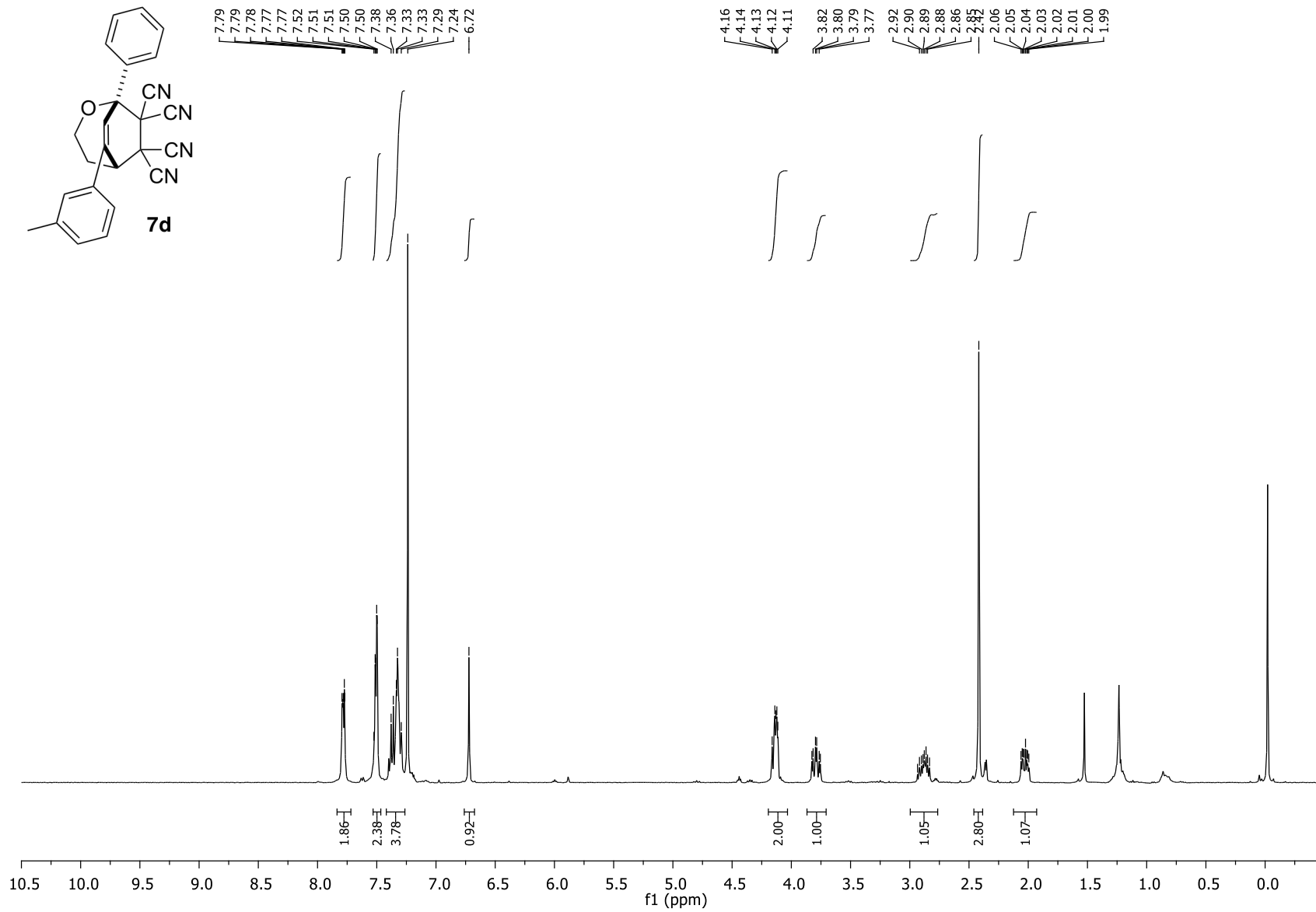


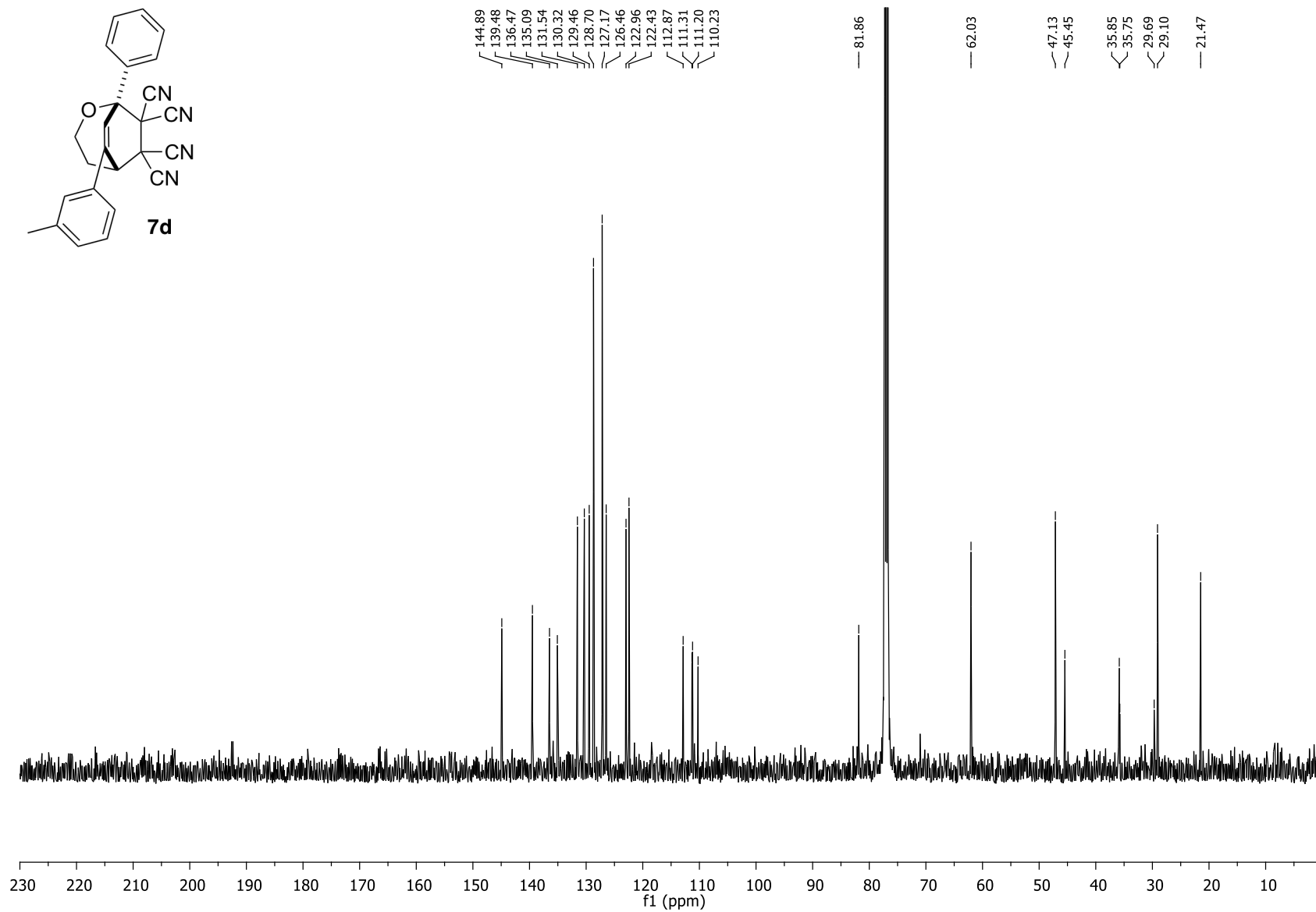
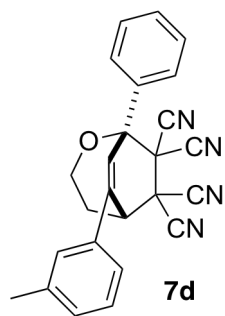


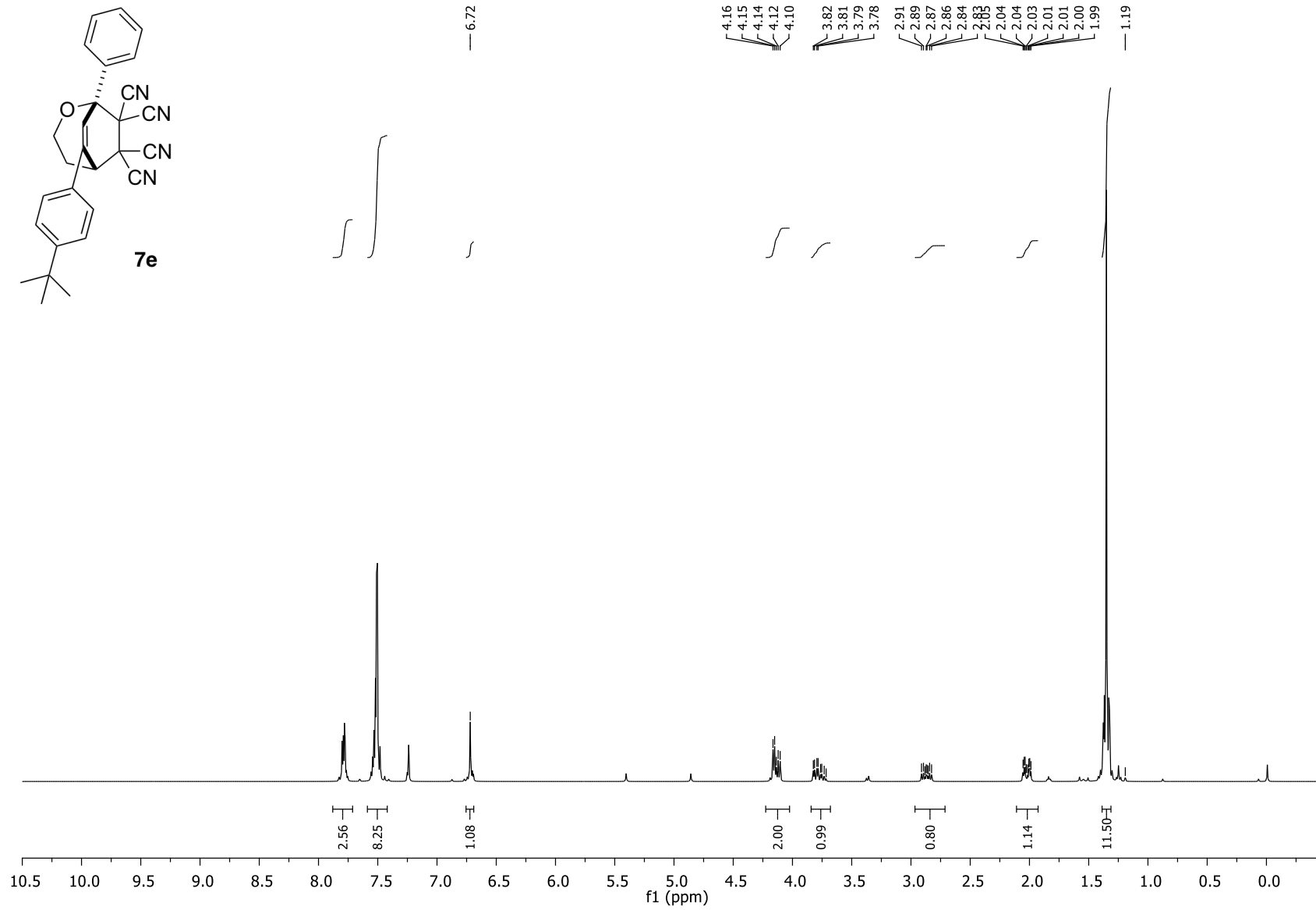
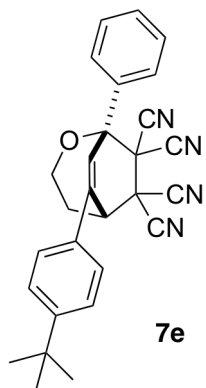


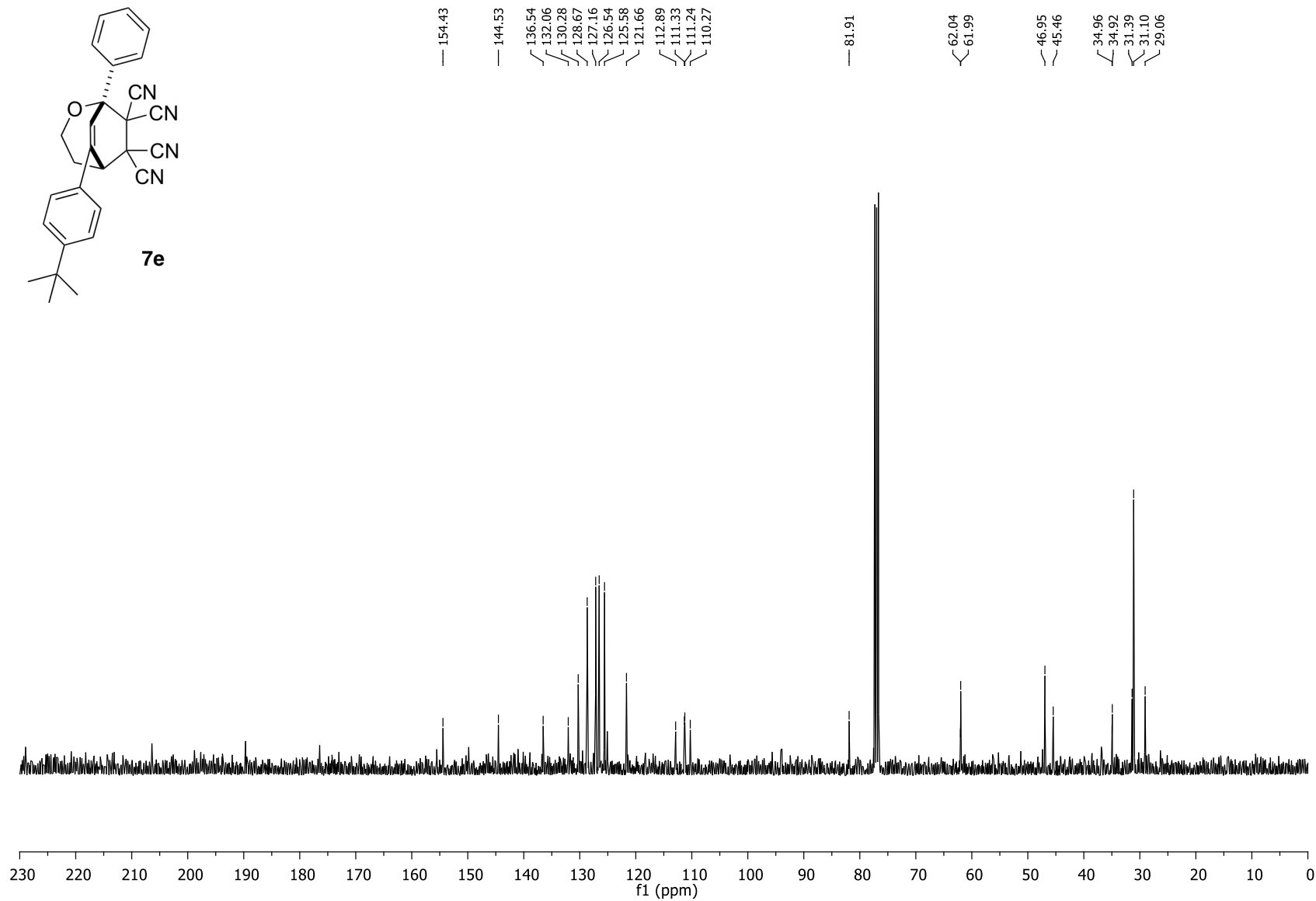
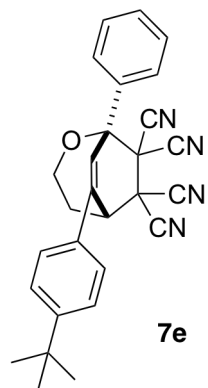


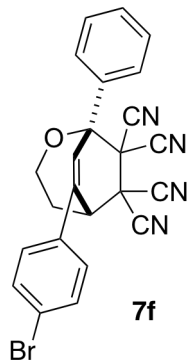










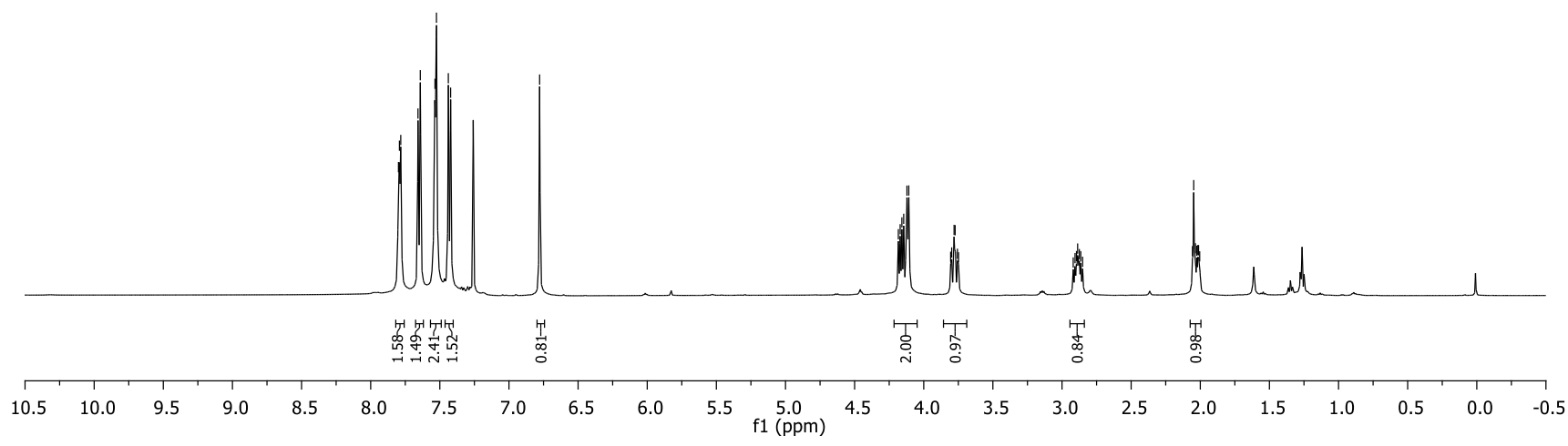


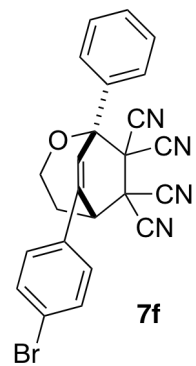
7.80
7.79
7.78
7.66
7.64
7.54
7.52
7.44
7.42
— 6.78

4.18
4.17
4.16
4.15
4.12
4.11

3.80
3.78
3.77
3.76

2.91
2.90
2.89
2.87
2.86
2.85
2.86
2.05
2.04
2.03
2.02
2.01
2.00





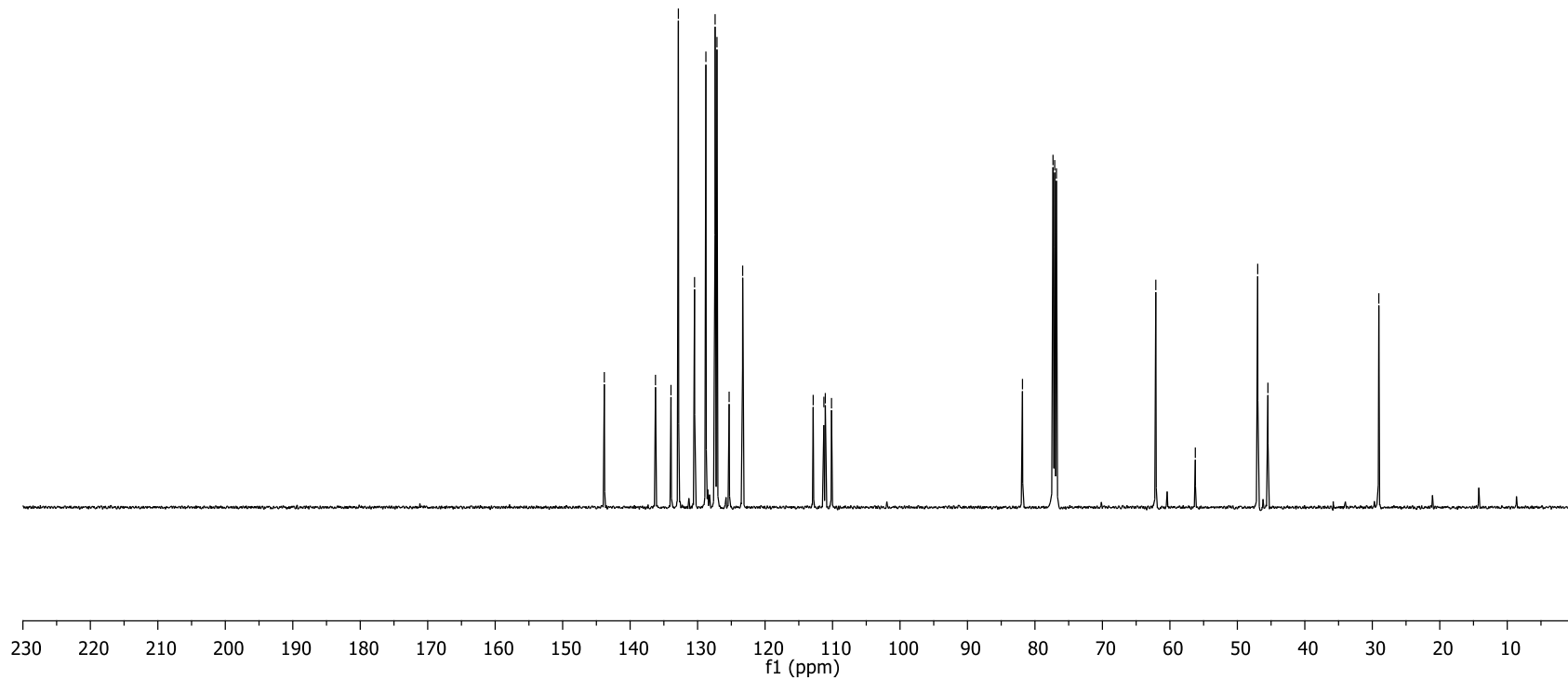
143.83
136.24
133.94
132.85
130.45
128.77
127.40
127.13
125.32
123.32
112.87
111.28
111.07
110.15

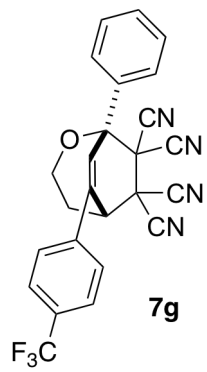
81.85
77.32
77.07
76.82

62.09
56.23

46.98
45.47

29.03





7.78
7.76
7.67
7.65
7.53
7.52
7.51
7.24
— 6.84

4.21
4.19
4.18
4.16
4.14
4.12
3.81
3.79
3.78
3.76
2.94
2.93
2.92
2.90
2.89
2.87
2.87
2.06
2.06
2.04
2.03
2.02
2.02
2.01

