

Supplementary Information

Electrochemical nickel-catalysed defluoroalkylation of gem-difluoroalkenes with alkyl halides

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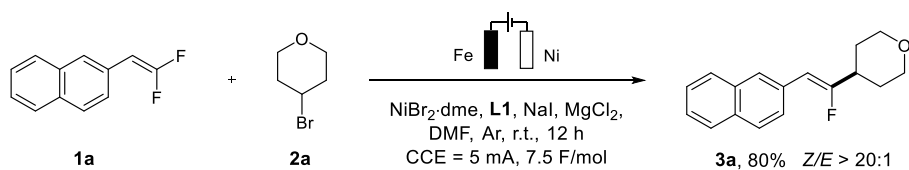
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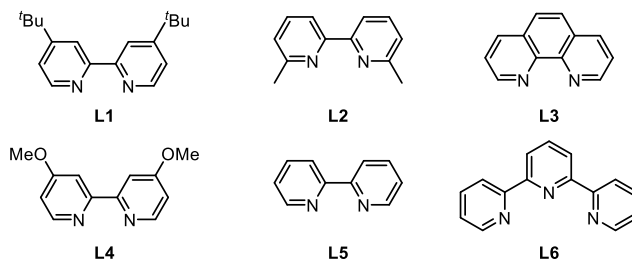
General Remarks

Catalytic reactions were carried out in undivided electrochemical cells (15 mL) using pre-dried glassware, if not noted otherwise. Solvents were obtained from commercial sources. All the starting materials were obtained from commercial sources or synthesized according to literature methods (>95% purity). Commercially available chemicals were obtained from *Bide Pharmatech Ltd, Tianjin Heowns OPDE Technologies* and *Shanghai Macklin Biochemical Co.* used as received unless otherwise stated. Iron plate electrodes (20.0 mm × 10.0 mm × 0.2 mm, 99.9%; obtained from Dingsheng scientific research metal materials, Hebei, China), Nickel foam electrodes (20.0 mm × 10.0 mm × 0.3 mm, 99.9%; obtained from Guangjiayuan electronic materials Jiangsu, China), were connected using stainless steel adapters. Electrocatalysis was conducted using an HSPY-36-03 potentiostat in constant current mode. Cyclic voltammetry studies were performed using a Shanghai Chenhua CHI760E workstation. Yields refer to isolated compounds, estimated to be >95% purity as determined by ¹H-NMR. Flash chromatography was performed using Silica gel (200-300 mesh) purchased from Qingdao Haiyang Chemical Co., China. NMR spectra were recorded on Bruker AVANCE AV 400 or 600 in the solvent indicated; chemical shifts are given in ppm relative to the residual solvent peak. The high-resolution mass spectrometry (HRMS) data were collected on a MicrOTOF mass spectrometer with ESI mass analyzer. All the synthesis methods of raw materials can be found in the corresponding literature.¹⁻⁵

Table S1: Optimization of the Reaction Conditions

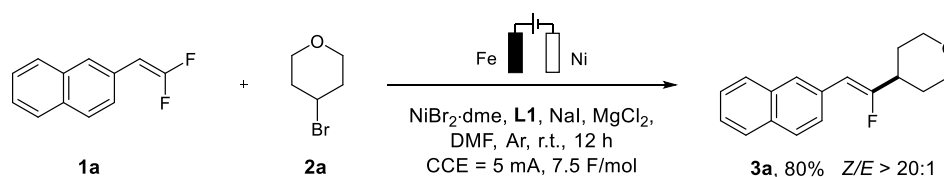
Entry	Variation	Yield %
1	none	80
2	w/o electricity	N.R.
3	w/o Nickel catalyst	19
4	w/o MgCl ₂	46
5	4 mA	30
6	6 mA	63
7	Mg(+) as anode	N.D.
8	Zn(+) as anode	trace
9	Cu(+) as anode	19
10	Al(+) as anode	29
11	Ni(+) as anode	16
12	Pt(+) as anode, DIPEA (2.0 equiv.)	3
13	GF(+) as anode, DIPEA (2.0 equiv.)	trace
14	CF(+) as anode, DIPEA (2.0 equiv.)	trace
15	Et ₄ NI instead of NaI	31
16	NH ₄ I instead of NaI	62
17	KI instead of NaI	trace
18	DMA instead of DMF	39
19	NiCl ₂ ·dme instead of NiBr ₂ ·dme	69
20	Ni(OTf) ₂ instead of NiBr ₂ ·dme	63
21	Ni(acac) ₂ instead of NiBr ₂ ·dme	55
22	NiI ₂ instead of NiBr ₂ ·dme	74
23	L2 instead of L1	Trace
24	L3 instead of L1	63
25	L4 instead of L1	33

26	L5 instead of L1	54
27	L6 instead of L1	18
28	AlCl ₃ instead of MgCl ₂	55
29	LiCl instead of MgCl ₂	62
30	LiBr instead of MgCl ₂	34
31	ZnCl ₂ instead of MgCl ₂	40



Reaction conditions: Fe plate anode, Ni foam cathode, **1a** (0.3 mmol), **2a** (0.9 mmol, 3.0 equiv.), NiBr₂·dme (10 mol%), 4,4'-di-*tert*-butyl-2,2'-bipyridine (**L1**, 20 mol%), MgCl₂ (2.0 equiv.), NaI (2.0 equiv.), dry DMF (5.0 mL), constant current = 5.0 mA, r.t., Ar, 12 h. Isolated yield. r.t. = room temperature. DIPEA = *N,N*-diisopropylethylamine. DMF = *N,N*-dimethylformamide. DMA = *N,N*-dimethylacetamide. CCE was constant current electrolysis, the amount of charge was 7.5 F/mol and the current density was 50 A/m². N.R. = no reaction. N.D. = not detected.

General Procedure for electrochemical nickel-catalyzed defluoroalkylation of *gem*-difluoroalkenes with alkyl halides



The electrocatalysis was carried out in an undivided cell with an iron plate anode (20.0 mm × 10.0 mm × 0.2 mm) and a Nickel foam cathode (20.0 mm × 10.0 mm × 0.3 mm). To a 15 mL pre-dried undivided electrochemical cell equipped with a magnetic bar were added **1a** (0.3 mmol, 1.0 equiv.), alkyl bromides (0.9 mmol, 3.0 equiv.), NaI (0.6 mmol, 2.0 equiv.), MgCl₂ (0.6 mmol, 2.0 equiv.), NiBr₂·dme (0.03 mmol, 0.1 equiv.) and **L1** (0.06 mmol, 0.2 equiv.). Then dry DMF (5 mL) was added after the reaction system was filled with argon. The electrocatalysis was performed at r.t. with a constant current of 5.0 mA and maintained for 12 h. The electrodes were washed with EtOAc (5 mL × 3) in an ultrasonic bath. Then, H₂O (20 mL) was added to the system, and the resulting mixture was extracted with EtOAc (50 mL × 2). The combined organic phase was dried with anhydrous Na₂SO₄, filtered, and concentrated in vacuo. The crude product was purified by column chromatography to furnish the desired product **3a**. However, some pseudohalides couldn't give good results, which were showed in the Figure **S1**.

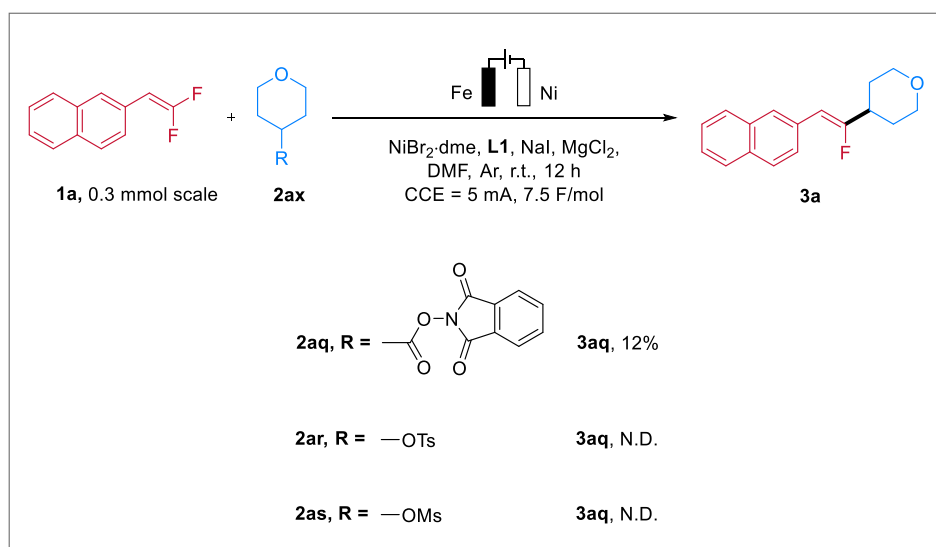


Figure **S1**. Failed results with pseudohalides

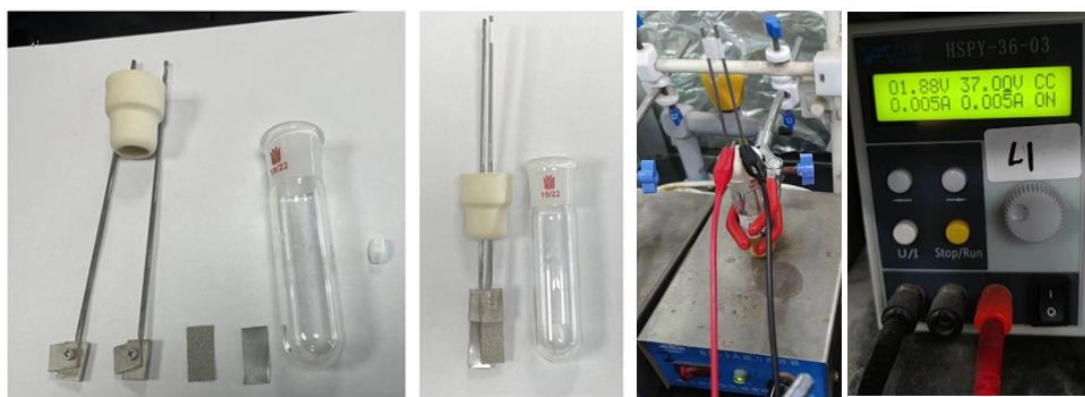
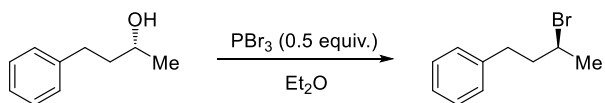


Figure S2. Electrolysis set-up

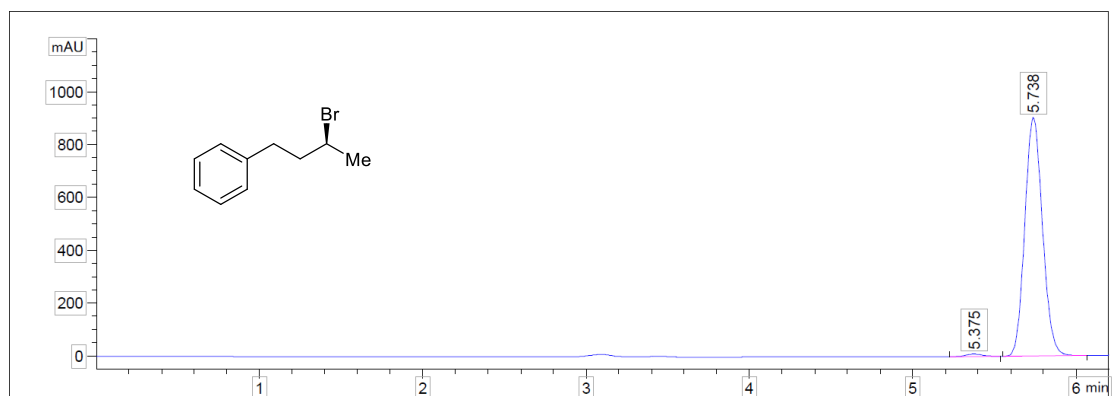
Mechanistic Experiments

Electrochemical nickel-catalyzed defluoroalkylation with gem-difluoroalkenes and alkyl halides

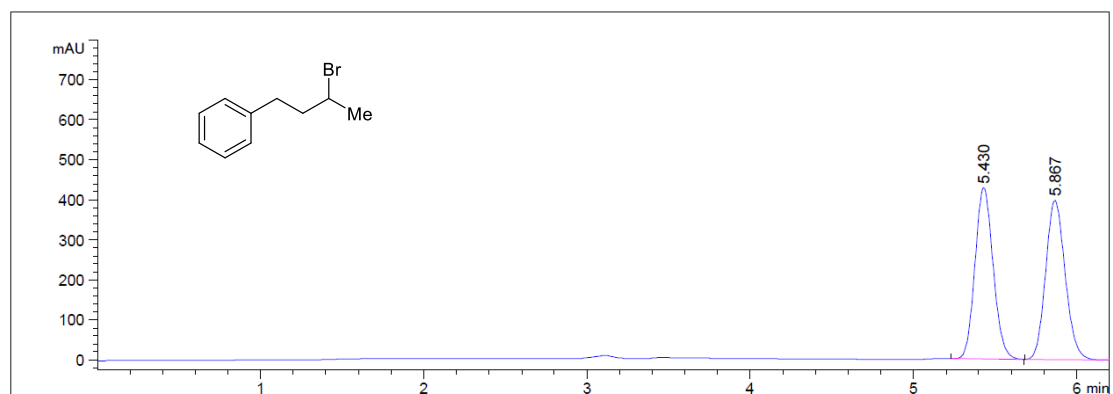
Reaction with chiral alkyl bromide



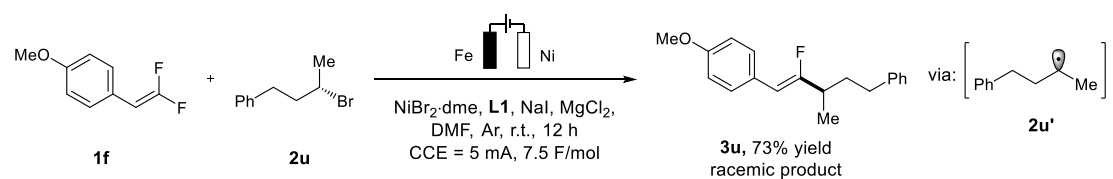
(*S*)-(3-bromobutyl)benzene was synthesized by brominating (*R*)-4-phenylbutan-2-ol according to reference^{5,6}. PBr_3 (2.5 mmol, 0.5 equiv.) was added dropwise to a 0 °C ice bath cooled solution of alcohol (5 mmol) in Et_2O (10 mL). The solution was then stirred at r.t. overnight. The reaction was quenched over ice water and stirred for 30 minutes. The aqueous layer was removed and extracted once with Et_2O . The organic layers were combined and washed sequentially with a saturated solution of NaHCO_3 , then brine, dried over MgSO_4 , filtered, and concentrated under reduced pressure. The crude product was purified by flash chromatography. ^1H NMR (400 MHz, Chloroform-*d*) δ 7.36 – 7.28 (m, 2H), 7.24 – 7.21 (m, 3H), 4.20 – 4.01 (m, 1H), 2.99 – 2.84 (m, 1H), 2.84 – 2.70 (m, 1H), 2.24 – 2.00 (m, 2H), 1.75 (d, $J = 6.7$ Hz, 3H). ^{13}C NMR (100 MHz, Chloroform-*d*) δ 141.1, 128.7, 128.6, 126.2, 51.1, 42.8, 34.1, 26.7. Physical and spectral data were in accordance with literature data^{5,6}. ee = 98%, HPLC analysis of the product: Chiralcel OJ-H column; 10% $^i\text{PrOH}$ in hexanes; 1.0 mL/min; retention times: 5.738 min (major), 5.375 min (minor).



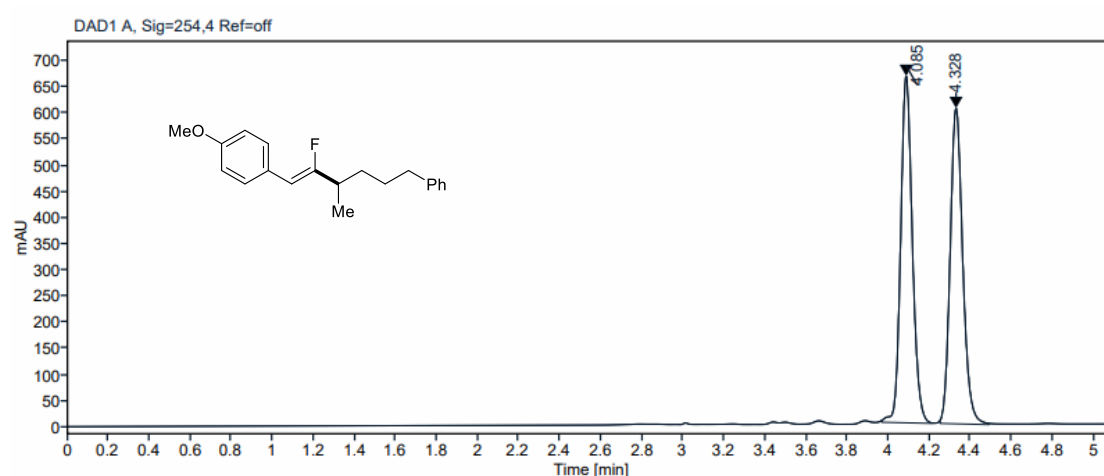
Peak #	Ret. Time	Area	Height	Area%
1	5.375	68.30441	9.58473	1.0012
2	5.738	6754.24121	902.48572	98.9988
Total		6822.54562	912.07045	100.000



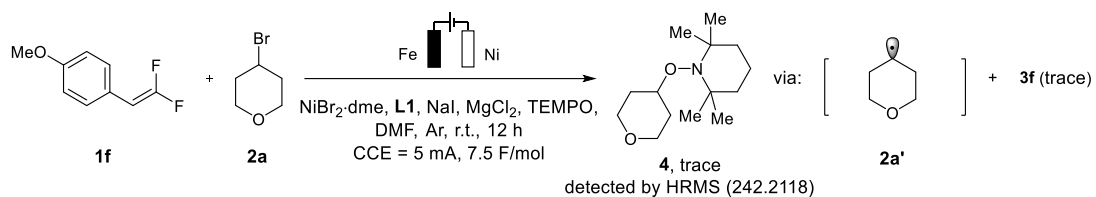
Peak #	Ret. Time	Area	Height	Area%
1	5.430	3303.96826	427.81207	49.9421
2	5.867	3311.63403	397.80164	50.0579
Total		6615.60229	825.61371	100.000



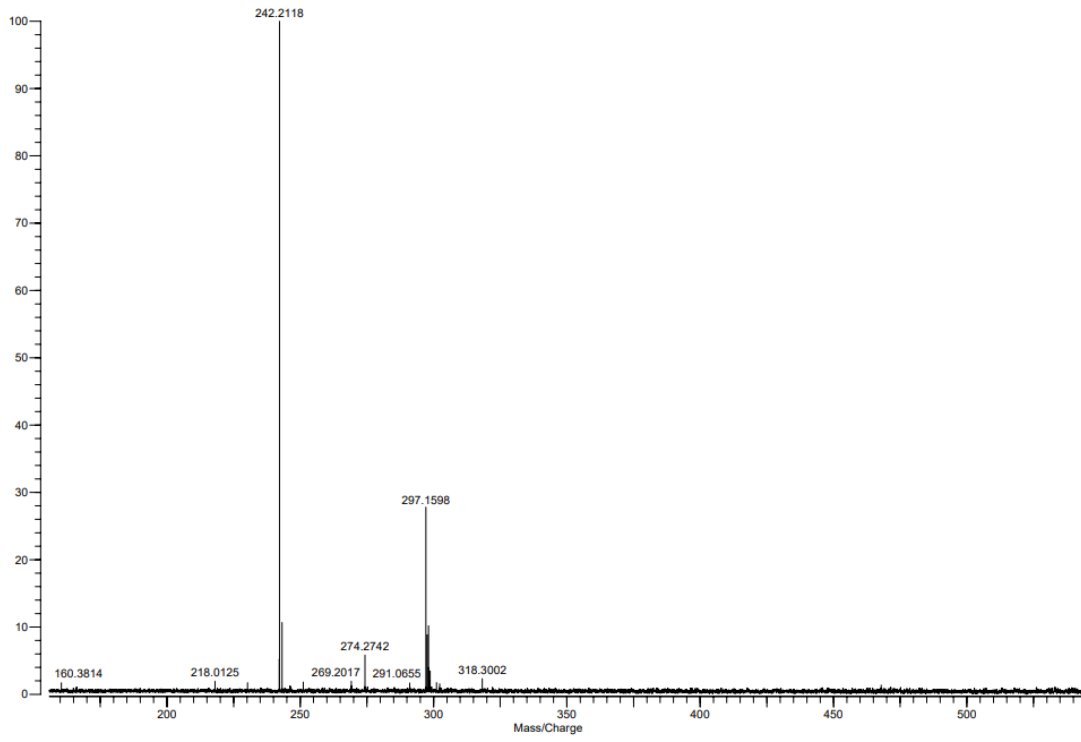
The electrocatalysis was carried out in an undivided cell with an iron plate anode (20.0 mm × 10.0 mm × 0.2 mm) and a Nickel foam cathode (20.0 mm × 10.0 mm × 0.3 mm). To a 15 mL pre-dried undivided electrochemical cell equipped with a magnetic bar were added **1f** (0.3 mmol, 1.0 equiv.), **2u** (0.9 mmol, 3.0 equiv.), NaI (0.6 mmol, 2.0 equiv.), MgCl₂ (0.6 mmol, 2.0 equiv.), NiBr₂·dme (0.03 mmol, 0.1 equiv.) and **L1** (0.06 mmol, 0.2 equiv.). And dry DMF (5 mL) was added after the reaction system was filled with argon. The electrocatalysis was performed at r.t. with a constant current of 5 mA and maintained for 12 h. The electrodes were washed with EtOAc (5 mL × 3) in an ultrasonic bath. H₂O (20 mL) was added to the system, and the resulting mixture was extracted with EtOAc (50 mL × 2). The combined organic phase was dried with anhydrous Na₂SO₄, filtered, and concentrated in vacuo. The crude product was purified by column chromatography to furnish the racemic product **3u** in 73% yield. ee = 0%, HPLC analysis of the product: Chiralcel AD-H column; 5% *i*PrOH in hexanes; 1.0 mL/min; retention times: 4.085 min, 4.328 min.



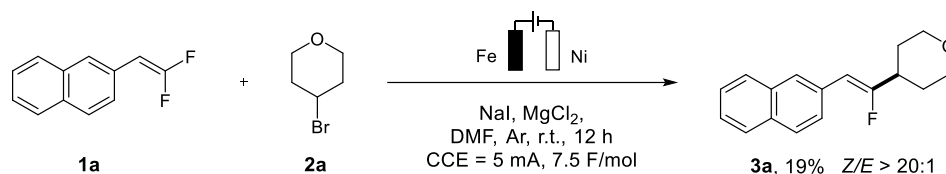
Reaction with TEMPO.



The electrocatalysis was carried out in an undivided cell with an iron plates anode (20.0 mm × 10.0 mm × 0.2 mm) and a Nickel foam cathode (20.0 mm × 10.0 mm × 0.3 mm). To a 15 mL pre-dried undivided electrochemical cell equipped with a magnetic bar were added **1f** (0.3 mmol, 1.0 equiv.), **2a** (0.9 mmol, 3.0 equiv.), TEMPO (0.9 mmol, 3.0 equiv.), $\text{NiBr}_2 \cdot \text{dme}$ (0.03 mmol, 0.1 equiv.), **L1** (0.06 mmol, 0.2 equiv.), NaI (0.03 mmol, 0.1 equiv.) and MgCl_2 (0.6 mmol, 2.0 equiv.). Then dry DMF (5 mL) was added after the reaction system was filled with argon. The electrocatalysis was performed at room temperature with a constant current of 5 mA maintained for 12 h. The electrodes were washed with EtOAc (5 mL × 3) in an ultrasonic bath. H_2O (20 mL) was added to the system, and the resulting mixture was extracted with EtOAc (50 mL × 2). The combined organic phase was dried with anhydrous Na_2SO_4 , filtered, and concentrated in vacuo. The crude product was purified by column chromatography to furnish the desired product. And **4** was detected by HRMS. m/z calc. for $\text{C}_{14}\text{H}_{27}\text{NO}_2$ $[\text{M}+\text{H}]^+$: 242.2120, found: 242.2118.



Electrochemical defluoroalkylation with *gem*-difluoroalkenes and alkyl halides



The electrocatalysis was carried out in an undivided cell with an iron plate anode (20.0 mm × 10.0 mm × 0.2 mm) and a Nickel foam cathode (20.0 mm × 10.0 mm × 0.3 mm). To a 15 mL pre-dried undivided electrochemical cell equipped with a magnetic bar were added **1a** (0.3 mmol, 1.0 equiv.), alkyl bromides (0.9 mmol, 3.0 equiv.), NaI (0.6 mmol, 2.0 equiv.), MgCl₂ (0.6 mmol, 2.0 equiv.). Then dry DMF (5 mL) was added after the reaction system was filled with argon. The electrocatalysis was performed at r.t. with a constant current of 5.0 mA and maintained for 12 h. The electrodes were washed with EtOAc (5 mL × 3) in an ultrasonic bath. Then, H₂O (20 mL) was added to the system, and the resulting mixture was extracted with EtOAc (50 mL × 2). The combined organic phase was dried with anhydrous Na₂SO₄, filtered, and concentrated in vacuo. The crude product was purified by column chromatography to furnish the desired product **3a**.

The possible mechanism was proposed according to corresponding literature.⁷ Firstly, the alkyl bromide was reduced at the cathode to form alkyl radical. And the radical addition process occurred between alkyl radical and *gem*-difluoroalkenes to generate **Int III**, after further reduction at the cathode to form Int IV, a β-F elimination could occur to afford target product.

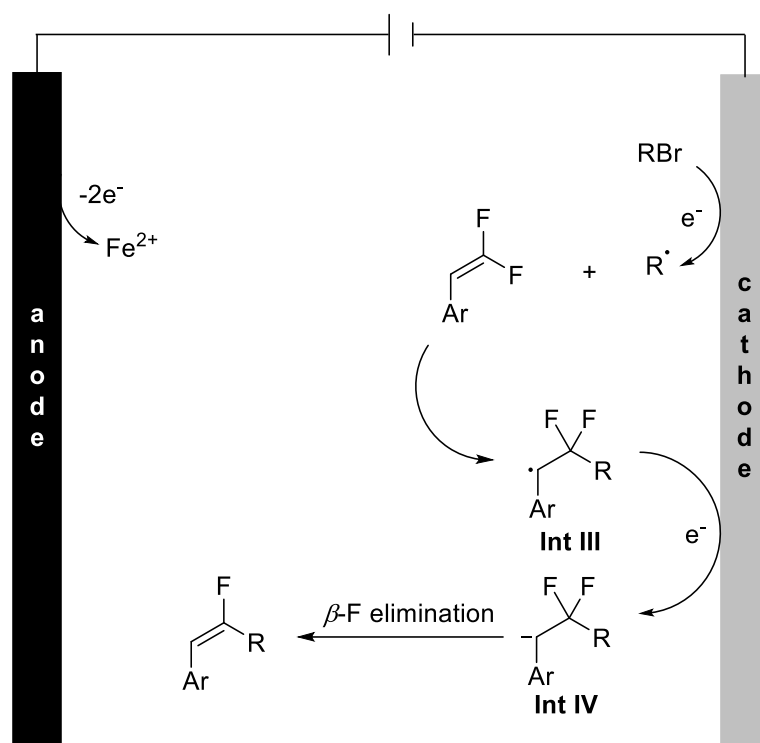


Figure S3. Proposal mechanisms

Cyclic Voltammetry

The cyclic voltammetry was carried out with a Shanghai Chenhua CHI760E workstation. A glassy-carbon (GC) electrode (5 mm-diameter, disk-electrode) was used as the working electrode, Pt wire was used as the counter electrode and an Ag/Ag⁺ electrode was used as the reference electrode. The measurements were carried out at a scan rate of 0.1 V s⁻¹ in DMF/TBAPF₆ (0.1 M) under reduction conditions. The operation temperature was 298 K.

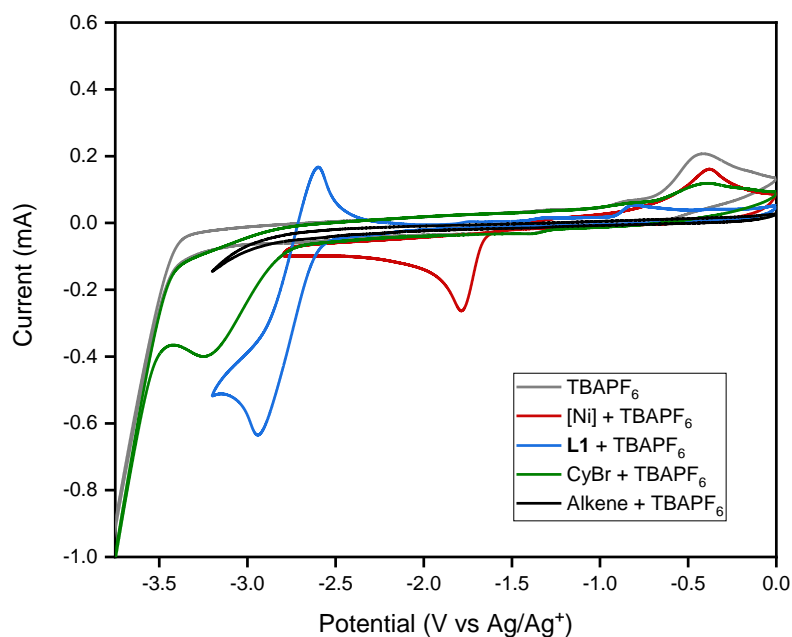


Figure S4. Cyclic voltammetry of TBAPF₆ (0.1 M), [Ni] (NiBr₂·dme, 0.005 M), L1 (4,4'-di-*tert*-butyl-2,2'-bipyridine, 0.01 M), CyBr (bromocyclohexane, **2w**, 0.01 M) and alkene (1-(2,2-difluorovinyl)-4-methoxybenzene, **1f**, 0.01 M) at 0.1 Vs⁻¹ in DMF (10.0 mL).

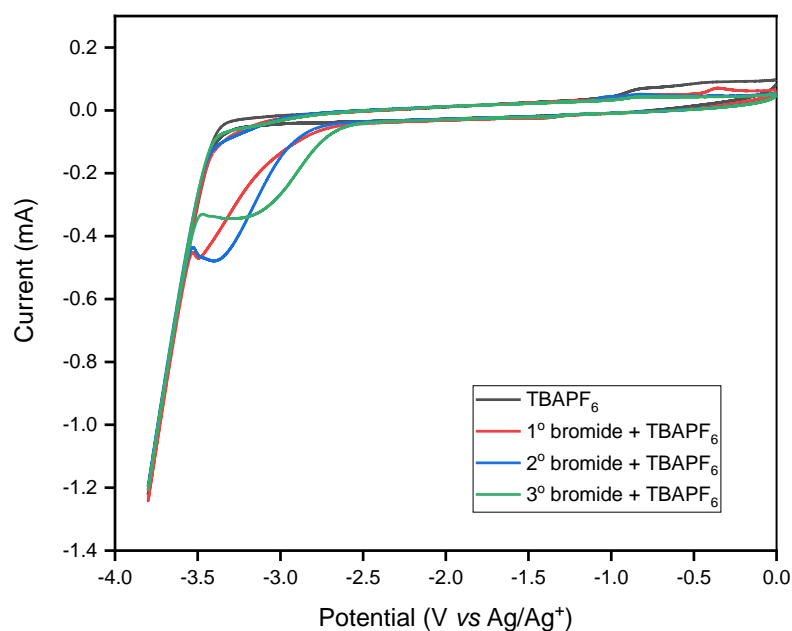


Figure S5. Cyclic voltammetry of TBAPF₆ (0.01 mM), 1° bromide ((4-bromobutoxy)benzene, **2ah**, 0.01 mM), 2° bromide (bromocyclohexane, **2w**, 0.01 M), 3° bromide (*t*BuBr, **2aa**, 0.01 M) at 0.1 Vs⁻¹ in DMF (10 mL).

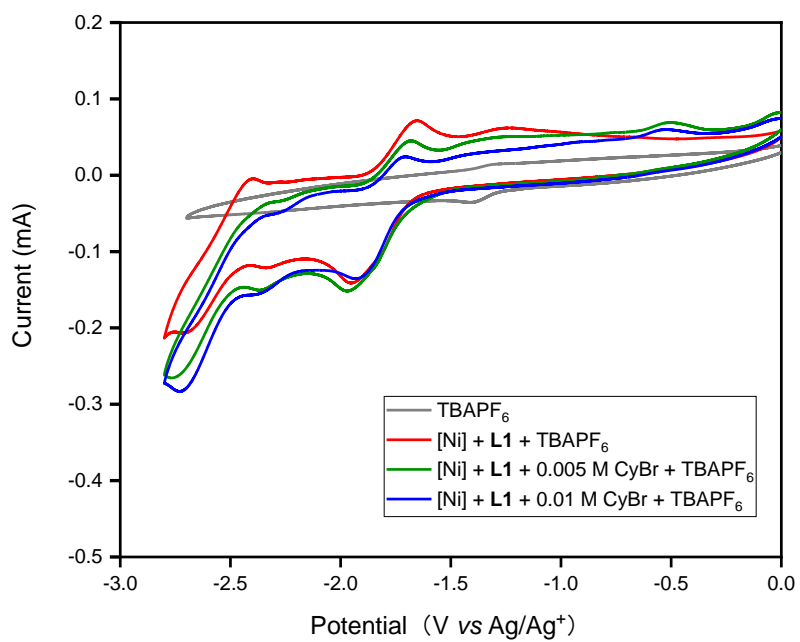


Figure S6. Cyclic voltammetry of TBAPF₆ (0.01 mM), NiBr₂·dme (0.005 M) + **L1** (0.01 M), NiBr₂·dme (0.005 M) + **L1** (0.01 M) + **2w** (0.005 M), NiBr₂·dme (0.005 M) + **L1** (0.01 M) + **2w** (0.01 M) at 0.1 Vs⁻¹ in DMF (10 mL).

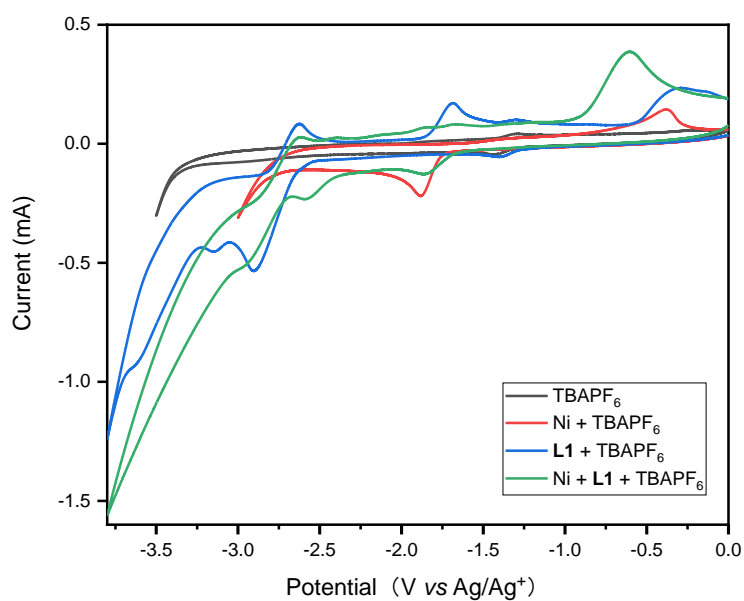


Figure S7: Cyclic voltammetry of TBAPF₆ (0.1 M), NiBr₂·dme (0.005 M), L1 (0.01 M), NiBr₂·dme + L1 (0.005 M + 0.01 M) at 0.1 Vs⁻¹ in DMF (10 mL).

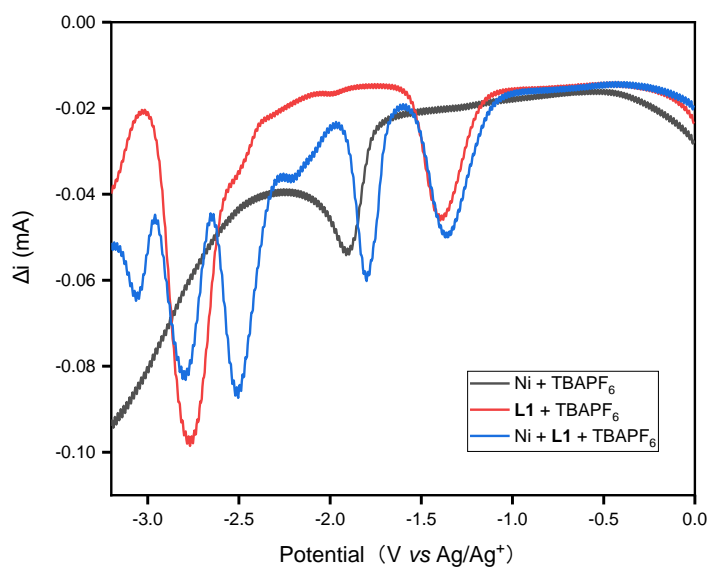
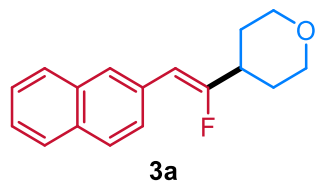


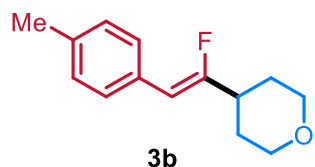
Figure S8: Square wave voltammetry of TBAPF₆ (0.1 M), NiBr₂·dme (0.005 M), L1 (0.01 M), NiBr₂·dme + L1 (0.005 M + 0.01 M) at 0.1 Vs⁻¹ in DMF (10 mL).

Characterization Data of Products



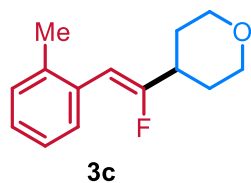
(Z)-4-(1-Fluoro-2-(naphthalen-2-yl)vinyl)tetrahydro-2H-pyran

Compound **3a** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 50:1) yielded **3a** (61.4 mg, 80%, *Z/E* > 20:1) as a light yellow solid. ^1H NMR (400 MHz, Chloroform-*d*) δ 7.91 (s, 1H), 7.80 – 7.78 (m, 3H), 7.66 – 7.64 (m, 1H), 7.48 – 7.43 (m, 2H), 5.63 (d, *J* = 40.3 Hz, 1H), 4.09 – 4.06 (m, 2H), 3.51 – 3.45 (m, 2H), 2.62 – 2.51 (m, 1H), 1.90 – 1.86 (m, 2H), 1.82 – 1.72 (m, 2H). ^{13}C NMR (150 MHz, Chloroform-*d*) δ 163.3 (d, *J* = 268.8 Hz), 133.6, 132.5, 131.2 (d, *J* = 2.1 Hz), 128.1 (d, *J* = 6.0 Hz), 127.7, 127.4 (d, *J* = 7.5 Hz), 126.8 (d, *J* = 7.5 Hz), 126.2, 126.0, 104.7 (d, *J* = 9.0 Hz), 67.7, 39.0 (d, *J* = 25.7 Hz), 29.9 (d, *J* = 3.0 Hz). ^{19}F NMR of (375 MHz, Chloroform-*d*): δ –106.38. HR-MS (ESI) *m/z* calc. for $\text{C}_{17}\text{H}_{18}\text{FO}^+$ [$\text{M}+\text{H}$] $^+$: 257.1336, found: 257.1341.



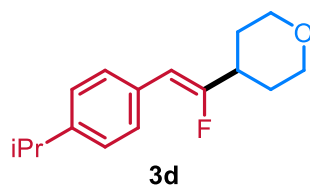
(Z)-4-(1-Fluoro-2-(p-tolyl)vinyl)tetrahydro-2H-pyran

Compound **3b** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 80:1) yielded **3b** (54.1 mg, 82%, *Z/E* > 20:1) as colorless liquid. ^1H (400 MHz, Chloroform-*d*) δ 7.39 – 7.37 (m, 2H), 7.14 – 7.12 (m, 2H), 5.43 (d, *J* = 40.6 Hz, 1H), 4.07 – 4.03 (m, 2H), 3.49 – 3.42 (m, 2H), 2.55 – 2.44 (m, 1H), 2.34 (s, 3H), 1.85 – 1.81 (m, 2H), 1.78 – 1.67 (m, 2H). ^{13}C NMR (100 MHz, Chloroform-*d*) δ 162.6 (d, *J* = 266.6 Hz), 136.8, 130.8 (d, *J* = 2.1 Hz), 129.3, 128.5 (d, *J* = 8.1 Hz), 104.4 (d, *J* = 10.1 Hz), 67.7, 38.9 (d, *J* = 26.2 Hz), 29.9 (d, *J* = 2.0 Hz), 21.3. ^{19}F NMR (375 MHz, Chloroform-*d*): δ –107.98. HR-MS (ESI) *m/z* calc. for $\text{C}_{14}\text{H}_{18}\text{FO}^+$ [$\text{M}+\text{H}$] $^+$: 221.1336, found: 221.1332.



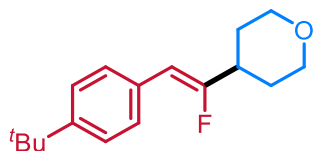
(Z)-4-(1-Fluoro-2-(o-tolyl)vinyl)tetrahydro-2H-pyran

Compound **3c** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 80:1) yielded **3c** (52.1 mg, 79%, *Z/E* > 20:1) as colorless liquid. ¹H (400 MHz, Chloroform-*d*) δ 7.61 – 7.59 (m, 1H), 7.19 – 7.13 (m, 3H), 5.60 (d, *J* = 39.3 Hz, 1H), 4.08 – 4.06 (m, 2H), 3.47 (t, *J* = 11.8 Hz, 2H), 2.57 – 2.51 (m, 1H), 2.30 (s, 3H), 1.87 – 1.85 (m, 2H), 1.79 – 1.72 (m, 2H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 162.9 (d, *J* = 265.8 Hz), 135.7, 132.1, 130.1, 129.3 (d, *J* = 9.0 Hz), 127.1, 126.0, 101.9 (d, *J* = 10.5 Hz), 67.6, 38.9 (d, *J* = 25.7 Hz), 30.0 (d, *J* = 3.0 Hz), 20.3. ¹⁹F NMR (375 MHz, Chloroform-*d*): δ –109.27. HR-MS (ESI) *m/z* calc. for C₁₄H₁₈FO⁺ [M+H]⁺: 221.1336, found: 221.1330.



(Z)-4-(1-Fluoro-2-(4-isopropylphenyl)vinyl)tetrahydro-2H-pyran

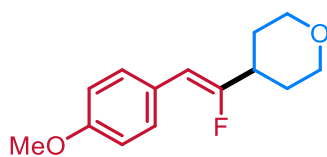
Compound **3d** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 80:1) yielded **3d** (55.8 mg, 75%, *Z/E* > 20:1) as light yellow oily liquid. ¹H (400 MHz, Chloroform-*d*) δ 7.43 – 7.41 (m, 2H), 7.20 – 7.18 (m, 2H), 5.45 (d, *J* = 40.6 Hz, 1H), 4.06 – 4.04 (m, 2H), 3.46 (t, *J* = 11.8 Hz, 2H), 2.93 – 2.86 (m, 1H), 2.56 – 2.45 (m, 1H), 1.85 – 1.81 (m, 2H), 1.79 – 1.67 (m, 2H), 1.25 (d, *J* = 6.9 Hz, 6H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 162.7 (d, *J* = 265.8 Hz), 147.8, 131.2, 128.5 (d, *J* = 7.6 Hz), 126.6, 104.3 (d, *J* = 9.1 Hz), 67.6, 38.9 (d, *J* = 25.7 Hz), 34.0, 29.9, 24.1. ¹⁹F NMR (375 MHz, Chloroform-*d*): δ –107.99. HR-MS (ESI) *m/z* calc. for C₁₆H₂₂FO⁺ [M+H]⁺: 249.1649, found: 249.1639.



3e

(Z)-4-(2-(4-(Tert-butyl)phenyl)-1-fluorovinyl)tetrahydro-2H-pyran

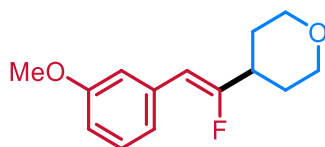
Compound **3e** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 80:1) yielded **3e** (59.0 mg, 75%, *Z/E* > 20:1) as light white solid. ¹H (400 MHz, Chloroform-*d*) δ 7.44 – 7.41 (m, 2H), 7.36 – 7.34 (m, 2H), 5.45 (d, *J* = 40.6 Hz, 1H), 4.07 – 4.04 (m, 2H), 3.49 – 3.42 (m, 2H), 2.55 – 2.46 (m, 1H), 1.84 – 1.81 (m, 2H), 1.77 – 1.67 (m, 2H), 1.30 (s, 9H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 162.8 (d, *J* = 265.8 Hz), 150.0, 130.8, 128.3 (d, *J* = 7.6 Hz), 125.5, 104.2 (d, *J* = 7.6 Hz), 67.6, 38.8 (d, *J* = 25.7 Hz), 34.7, 31.4 (d, *J* = 7.6 Hz), 29.9. ¹⁹F NMR (375 MHz, Chloroform-*d*): δ –107.85. HR-MS (ESI) *m/z* calc. for C₁₇H₂₄FO⁺ [M+H]⁺ : 263.1806, found: 263.1816.



3f

(Z)-4-(1-Fluoro-2-(4-methoxyphenyl)vinyl)tetrahydro-2H-pyran

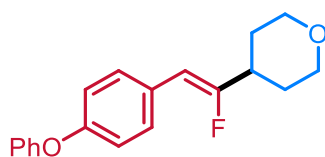
Compound **3f** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 50:1) yielded **3f** (55.2 mg, 78%, *Z/E* = 13:1) as colorless liquid. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.43 – 7.41 (m, 2H), 6.87 – 6.85 (m, 2H), 5.40 (d, *J* = 40.6 Hz, 1H), 4.06 – 3.99 (m, 2H), 3.81 (s, 3H), 3.48 – 3.36 (m, 2H), 2.53 – 2.44 (m, 1H), 1.84 – 1.81 (m, 2H), 1.77 – 1.66 (m, 2H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 161.9 (d, *J* = 264.3 Hz), 158.5, 129.8 (d, *J* = 7.6 Hz), 126.4, 114.0, 103.9 (d, *J* = 9.1 Hz) 67.7, 55.4, 38.8 (d, *J* = 25.7 Hz), 29.9 (d, *J* = 3.0 Hz). ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –106.11. HR-MS (ESI) *m/z* calc. for C₁₄H₁₈FO₂⁺ [M+H]⁺ : 237.1285, found: 237.1276.



3g

(Z)-4-(1-Fluoro-2-(3-methoxyphenyl)vinyl)tetrahydro-2H-pyran

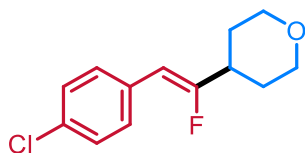
Compound **3g** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 50:1) yielded **3g** (55.2 mg, 75%, *Z/E* = 13:1) as colorless liquid. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.31 – 7.28 (m, 1H), 7.12 – 7.08 (m, 2H), 6.84 – 6.82 (m, 1H), 5.49 (d, *J* = 40.1 Hz, 1H), 4.12 – 4.08 (m, 2H), 3.86 (s, 3H), 3.53 – 3.47 (m, 2H), 2.60 – 2.51 (m, 1H), 1.89 – 1.86 (m, 2H), 1.83 – 1.72 (m, 2H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 163.4 (d, *J* = 267.3 Hz) 159.7, 134.9, 129.5, 121.2 (d, *J* = 6.0 Hz), 113.9 (d, *J* = 7.5 Hz), 112.9, 104.5 (d, *J* = 9.0 Hz), 67.6, 55.3, 39.0 (d, *J* = 25.7 Hz), 29.9 (d, *J* = 1.5 Hz). ¹⁹F NMR (375 MHz, Chloroform-*d*) δ – 106.11. HR-MS (ESI) *m/z* calc. for C₁₄H₁₈FO₂⁺ [M+H]⁺ : 237.1285, found: 237.1295.



3h

(Z)-4-(1-Fluoro-2-(4-phenoxyphenyl)vinyl)tetrahydro-2H-pyran

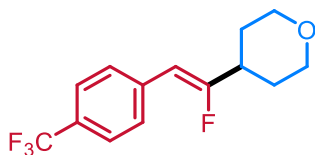
Compound **3h** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 80:1) yielded **3h** (63.5 mg, 71%, *Z/E* > 20:1) as white solid. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.35 – 7.08 (m, 6H), 7.02 – 7.00 (m, 2H), 6.88 – 6.86 (m, 1H), 5.43 (d, *J* = 39.7 Hz, 1H), 4.06 – 4.02 (m, 2H), 3.44 (t, *J* = 11.8 Hz, 2H), 2.53 – 2.44 (m, 1H), 1.83 – 1.80 (m, 2H), 1.76 – 1.65 (m, 2H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 163.7 (d, *J* = 267.0 Hz), 157.4, 157.3, 135.4, 129.9, 129.8, 123.7 (d, *J* = 7.5 Hz) 123.3, 119.1 (d, *J* = 7.5 Hz), 118.9, 117.7, 104.1 (d, *J* = 7.5 Hz), 67.6, 38.9 (d, *J* = 25.5 Hz), 29.8 (d, *J* = 3.0 Hz). ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –105.35. HR-MS (ESI) *m/z* calc. for C₁₉H₁₉FNaO₂⁺ [M+Na]⁺ : 321.1261, found: 321.1261.



3i

(Z)-4-(2-(4-Chlorophenyl)-1-fluorovinyl)tetrahydro-2H-pyran

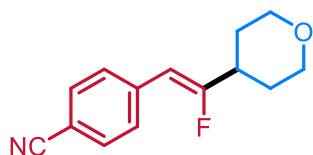
Compound **3i** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 50:1) yielded **3i** (52.6 mg, 73%, *Z/E* > 20:1) as white solid. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.42 – 7.39 (m, 2H), 7.30 – 7.27 (m, 2H), 5.43 (d, *J* = 39.8 Hz, 1H), 4.07 – 4.03 (m, 2H), 3.48 – 3.42 (m, 2H), 2.56 – 2.45 (m, 1H), 1.84 – 1.81 (m, 2H), 1.77 – 1.67 (m, 2H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 163.6 (d, *J* = 265.5 Hz), 132.6 (d, *J* = 3.0 Hz), 132.1, 129.8 (d, *J* = 7.5 Hz), 128.7, 103.6 (d, *J* = 9.0 Hz), 67.6, 38.8 (d, *J* = 27.2 Hz), 29.8 (d, *J* = 1.5 Hz). ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –105.99. HR-MS (ESI) *m/z* calc. for C₁₃H₁₅ClFO⁺ [M+H]⁺ : 241.0790, found: 241.0785



3j

(Z)-4-(1-Fluoro-2-(4-(trifluoromethyl)phenyl)vinyl)tetrahydro-2H-pyran

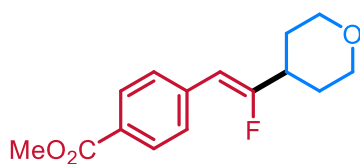
Compound **3j** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 50:1) yielded **3j** (50.1 mg, 61%, *Z/E* > 20:1) as white solid. ¹H NMR (600 MHz, Chloroform-*d*) δ 7.61 – 7.51 (m, 4H), 5.52 (d, *J* = 39.4 Hz, 1H), 4.08 – 4.05 (m, 2H), 3.48 – 3.44 (m, 2H), 2.55 – 2.52 (m, 1H), 1.86 – 1.83 (m, 2H), 1.77 – 1.70 (m, 2H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 164.9 (d, *J* = 270.3 Hz), 137.2, 128.7 (d, *J* = 7.5 Hz), 125.5 (q, *J* = 4.5 Hz), 124.3 (q, *J* = 271.2 Hz), 103.6 (d, *J* = 7.5 Hz), 67.5, 39.0 (d, *J* = 25.7 Hz), 29.7 (d, *J* = 3.0 Hz). ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –62.53, –103.54. HR-MS (ESI) *m/z* calc. for C₁₄H₁₅F₄O⁺ [M+H]⁺ : 275.1054, found: 275.1057.



3k

(Z)-4-(2-Fluoro-2-(tetrahydro-2H-pyran-4-yl)vinyl)benzonitrile

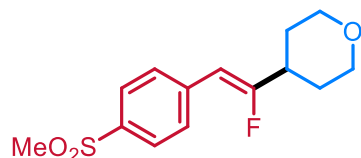
Compound **3k** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 30:1) yielded **3k** (35.3 mg, 51%, *Z/E* = 17:1) as white solid. ¹H NMR (600 MHz, Chloroform-*d*) δ 7.60 – 7.59 (m, 2H), 7.56 – 7.55 (m, 2H), 5.51 (d, *J* = 39.0 Hz, 1H), 4.08 – 4.05 (m, 2H), 3.46 (t, *J* = 11.9 Hz, 2H), 2.58 – 2.51 (m, 1H), 1.86 – 1.83 (m, 2H), 1.77 – 1.70 (m, 2H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 165.8 (d, *J* = 271.8 Hz), 138.4, 132.4, 129.0 (d, *J* = 7.5 Hz), 119.1, 110.3 (d, *J* = 3.0 Hz), 103.7 (d, *J* = 7.5 Hz), 67.5, 39.0 (d, *J* = 25.7 Hz), 29.7 (d, *J* = 3.0 Hz). ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –101.18. HR-MS (ESI) *m/z* calc. for C₁₄H₁₅FNO⁺ [M+H]⁺ : 232.1132, found: 232.1134.



3l

Methyl (Z)-4-(2-fluoro-2-(tetrahydro-2H-pyran-4-yl)vinyl)benzoate

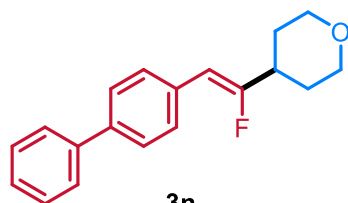
Compound **3l** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 30:1) yielded **3l** (23.8 mg, 30%, *Z/E* > 20:1) as white solid. ¹H NMR (400 MHz, Chloroform-*d*) δ 8.00 – 7.97 (m, 2H), 7.55 – 7.52 (m, 2H), 5.52 (d, *J* = 39.7 Hz, 1H), 4.08 – 4.04 (m, 2H), 3.91 (s, 3H), 3.49 – 3.43 (m, 2H), 2.59 – 2.48 (m, 1H), 1.87 – 1.82 (m, 2H), 1.79 – 1.68 (m, 2H). ¹³C NMR (100 MHz, Chloroform-*d*) δ 167.0, 164.9 (d, *J* = 271.7 Hz), 138.3 (d, *J* = 3.0 Hz), 129.9, 128.4 (d, *J* = 7.5 Hz), 104.1 (d, *J* = 8.9 Hz), 67.6, 52.2, 39.0 (d, *J* = 26.3 Hz), 30.2, 29.8 (d, *J* = 2.0 Hz). ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –102.86. HR-MS (ESI) *m/z* calc. for C₁₅H₁₈FO₃⁺ [M+H]⁺ : 265.1234, found: 265.1235.



3m

(Z)-4-(1-Fluoro-2-(4-(methylsulfonyl)phenyl)vinyl)tetrahydro-2H-pyran

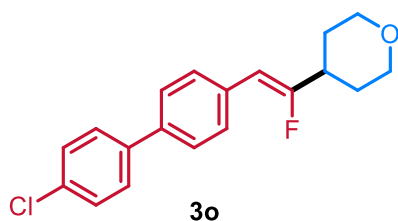
Compound **3m** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 30:1) yielded **3m** (51.1 mg, 60%, *Z/E* > 20:1) as white solid. ^1H NMR (400 MHz, Chloroform-*d*) δ 7.88 – 7.86 (m, 2H), 7.65 – 7.63 (m, 2H), 5.55 (d, *J* = 39.0 Hz, 1H), 4.07 – 4.04 (m, 2H), 3.49 – 3.43 (m, 2H), 3.04 (s, 3H), 2.61 – 2.50 (m, 1H), 1.87 – 1.82 (m, 2H), 1.79 – 1.68 (m, 2H). ^{13}C NMR (100 MHz, Chloroform-*d*) δ 165.9 (d, *J* = 273.7 Hz), 139.2 (d, *J* = 2.0 Hz), 138.3 (d, *J* = 3.0 Hz), 129.2 (d, *J* = 8.0 Hz), 127.7, 103.4 (d, *J* = 8.0 Hz), 67.5, 44.7, 39.0 (d, *J* = 25.3 Hz), 29.6 (d, *J* = 3.0 Hz). ^{19}F NMR (375 MHz, Chloroform-*d*) δ –101.27. HR-MS (ESI) *m/z* calc. for $\text{C}_{14}\text{H}_{17}\text{FNaO}_3\text{S}^+$ [*M*+*Na*] $^+$: 307.0775, found: 307.0765.



3n

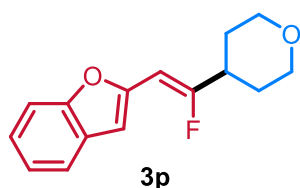
(Z)-4-(2-([1,1'-Biphenyl]-4-yl)-1-fluorovinyl)tetrahydro-2H-pyran

Compound **3n** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 50:1) yielded **3n** (55.0 mg, 65%, *Z/E* > 20:1) as light yellow oily liquid. ^1H NMR (600 MHz, Chloroform-*d*) δ 7.61 – 7.56 (m, 6H), 7.45 – 7.42 (m, 2H), 7.35 – 7.32 (m, 1H), 5.52 (d, *J* = 40.3 Hz, 1H), 4.07 – 4.05 (m, 2H), 3.48 – 3.44 (m, 2H), 2.58 – 2.50 (m, 1H), 1.86 – 1.84 (m, 2H), 1.78 – 1.71 (m, 2H). ^{13}C NMR (150 MHz, Chloroform-*d*) δ 163.4 (d, *J* = 265.5 Hz), 140.8, 139.7, 132.7, 129.0 (d, *J* = 7.5 Hz), 128.9, 127.4, 127.2, 127.1, 104.2 (d, *J* = 7.5 Hz), 67.6, 38.9 (d, *J* = 25.5 Hz), 29.9 (d, *J* = 3.0 Hz). ^{19}F NMR (375 MHz, Chloroform-*d*) δ –106.34. HR-MS (ESI) *m/z* calc. for $\text{C}_{19}\text{H}_{19}\text{NaFO}^+$ [*M*+*Na*] $^+$: 305.1312, found: 305.1308.



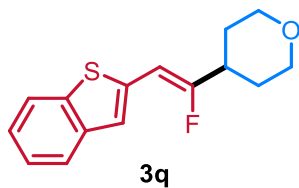
(Z)-4-(2-(4'-Chloro-[1,1'-biphenyl]-4-yl)-1-fluorovinyl)tetrahydro-2H-pyran

Compound **3o** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 50:1) yielded **3o** (66.4 mg, 70%, *Z/E* > 20:1) as white solid. ¹H NMR (600 MHz, Chloroform-*d*) δ 7.57 – 7.51 (m, 6H), 7.41 – 7.39 (m, 2H), 5.51 (d, *J* = 40.2 Hz, 1H), 4.08 – 4.05 (m, 2H), 3.50 – 3.34 (m, 2H), 2.58 – 2.49 (m, 1H), 1.87 – 1.84 (m, 2H), 1.80 – 1.70 (m, 2H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 163.6 (d, *J* = 267.3 Hz), 139.3, 138.4, 133.5, 133.1, 129.1, 129.1, 128.3, 127.1, 104.1 (d, *J* = 9.0 Hz), 67.6, 39.0 (d, *J* = 27.2 Hz), 29.9 (d, *J* = 3.0 Hz). ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –105.97. HR-MS (ESI) *m/z* calc. for C₁₉H₁₉ClFO⁺ [M+H]⁺ : 317.1103, found: 317.1096.



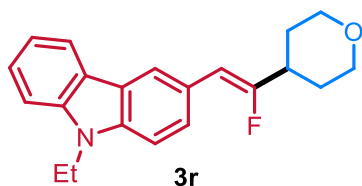
(Z)-2-(2-Fluoro-2-(tetrahydro-2H-pyran-4-yl)vinyl)benzofuran

Compound **3p** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 80:1) yielded **3p** (26.6 mg, 36%, *Z/E* > 50:1) as white solid. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.54 – 7.53 (m, 1H), 7.44 – 7.42 (m, 1H), 7.27 – 7.18 (m, 2H), 6.87 (s, 1H), 5.69 (d, *J* = 38.4 Hz, 1H), 4.08 – 4.04 (m, 2H), 3.50 – 3.43 (m, 2H), 2.63 – 2.52 (m, 1H), 1.88 – 1.84 (m, 2H), 1.78 – 1.68 (m, 2H). ¹³C NMR (100 MHz, Chloroform-*d*) δ 164.5 (d, *J* = 272.7 Hz), 154.0, 150.7 (d, *J* = 2.0 Hz), 129.3, 124.2, 123.0, 120.9, 111.0, 105.5 (d, *J* = 11.1 Hz), 95.9 (d, *J* = 11.1 Hz), 67.5, 38.5 (d, *J* = 25.3 Hz), 29.6 (d, *J* = 3.0 Hz). ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –97.78. HR-MS (ESI) *m/z* calc. for C₁₅H₁₆FO₂⁺ [M+H]⁺ : 247.1129, found: 247.1122.



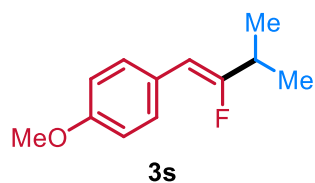
(Z)-4-(2-(Benzo[b]thiophen-2-yl)-1-fluorovinyl)tetrahydro-2H-pyran

Compound **3q** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 80:1) yielded **3q** (27.5 mg, 35%, *Z/E* > 50:1) as white solid. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.80 – 7.78 (m, 1H), 7.72 – 7.70 (m, 1H), 7.35 – 7.26 (m, 2H), 7.23 (s, 1H), 5.87 (d, *J* = 38.9 Hz, 1H), 4.09 – 4.04 (m, 2H), 3.50 – 3.43 (m, 2H), 2.63 – 2.52 (m, 1H), 1.88 – 1.84 (m, 2H), 1.79 – 1.69 (m, 2H). ¹³C NMR (100 MHz, Chloroform-*d*) δ 163.5 (d, *J* = 270.7 Hz), 140.1 (d, *J* = 8.0 Hz), 139.4, 136.1 (d, *J* = 4.0 Hz), 124.5, 124.4, 123.3 (d, *J* = 3.0 Hz), 122.7 (d, *J* = 4.0 Hz), 122.2, 99.9 (d, *J* = 12.1 Hz), 67.5, 38.4 (d, *J* = 25.0 Hz), 29.7 (d, *J* = 2.0 Hz). ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –101.56. HR-MS (ESI) *m/z* calc. for C₁₅H₁₆FOS⁺ [M+H]⁺ : 263.0906, found: 263.0910.



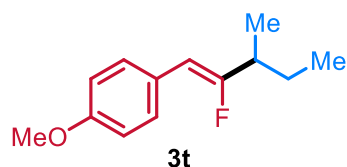
(Z)-9-Ethyl-3-(2-fluoro-2-(tetrahydro-2H-pyran-4-yl)vinyl)-9H-carbazole

Compound **3r** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 50:1) yielded **3r** (65.9 mg, 68%, *Z/E* = 9:1) as white solid. ¹H NMR (400 MHz, Chloroform-*d*) δ 8.25 (s, 1H), 8.11 – 8.09 (m, 1H), 7.63 – 7.60 (m, 1H), 7.49 – 7.45 (m, 1H), 7.41 – 7.39 (m, 1H), 7.37 – 7.34 (m, 1H), 7.25 – 7.21 (m, 1H), 5.65 (d, *J* = 40.9 Hz, 1H), 4.36 (q, *J* = 7.2 Hz, 2H), 4.10 – 4.06 (m, 2H), 3.52 – 3.46 (m, 2H), 2.61 – 2.51 (m, 1H), 1.91 – 1.87 (m, 2H), 1.84 – 1.74 (m, 2H), 1.43 (t, *J* = 7.2 Hz, 3H). ¹³C NMR (100 MHz, Chloroform-*d*) δ 161.5 (d, *J* = 264.6 Hz), 140.4, 139.1, 126.7 (d, *J* = 40.9 Hz), 125.9, 124.6, 123.2 (d, *J* = 9.0 Hz), 120.6, 120.5, 119.04, 108.7, 108.5, 105.0 (d, *J* = 9.1 Hz), 67.8, 39.0 (d, *J* = 26.3 Hz), 37.7, 30.1 (d, *J* = 2.0 Hz), 14.0. ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –110.55. HR-MS (ESI) *m/z* calc. for C₂₁H₂₃FNO⁺ [M+H]⁺ : 324.1758, found: 324.1753.



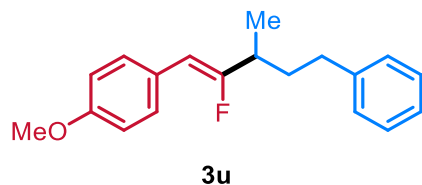
(Z)-1-(2-Fluoro-3-methylbut-1-en-1-yl)-4-methoxybenzene

Compound **3s** was prepared following the general procedure, purification by column chromatography on silica gel (PE) yielded **3s** (37.8 mg, 65%, *Z/E* > 20:1) as colourless liquid. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.43 – 7.41 (m, 2H), 6.87 – 6.84 (m, 2H), 5.40 (d, *J* = 40.6 Hz, 1H), 3.81 (s, 3H), 2.60 – 2.50 (m, 1H), 1.18 (d, *J* = 6.9 Hz, 6H). ¹³C NMR (100 MHz, Chloroform-*d*) δ 164.6 (d, *J* = 266.6 Hz), 158.3 (d, *J* = 2.0 Hz), 129.7 (d, *J* = 7.5 Hz), 126.8 (d, *J* = 3.0 Hz), 114.0, 102.8 (d, *J* = 9.1 Hz), 55.4, 32.1 (d, *J* = 25.3 Hz), 20.0 (d, *J* = 2.0 Hz). ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –109.40. HR-MS (ESI) *m/z* calc. for C₁₂H₁₅NaFO⁺ [M+Na]⁺ : 217.0999, found: 217.1004. The analytical data corresponds with those reported in the literature.⁸



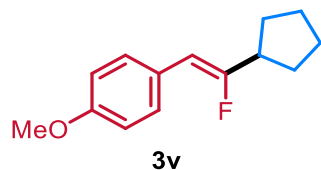
(Z)-1-(2-Fluoro-3-methylpent-1-en-1-yl)-4-methoxybenzene

Compound **3t** was prepared following the general procedure, purification by column chromatography on silica gel (PE) yielded **3t** (43.1 mg, 69%, *Z/E* = 16:1) as colorless liquid. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.43 – 7.41 (m, 2H), 6.87 – 6.84 (m, 2H), 5.40 (d, *J* = 40.5 Hz, 1H), 3.81 (s, 3H), 2.35 – 2.21 (m, 1H), 1.68 – 1.57 (m, 1H), 1.50 – 1.39 (m, 1H), 1.16 (d, *J* = 6.9 Hz, 3H), 0.94 (t, *J* = 7.4 Hz, 3H). ¹³C NMR (100 MHz, Chloroform-*d*) δ 163.1 (d, *J* = 267.7 Hz), 158.3 (d, *J* = 3.0 Hz), 129.7 (d, *J* = 8.1 Hz), 126.9, 113.9, 104.3 (d, *J* = 9.1 Hz), 55.4, 39.4 (d, *J* = 25.3 Hz), 26.9, 17.7 (d, *J* = 3.0 Hz), 11.8. ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –112.09. HR-MS (ESI) *m/z* calc. for C₁₃H₁₇NaFO⁺ [M+Na]⁺ : 231.1156, found: 231.1158.



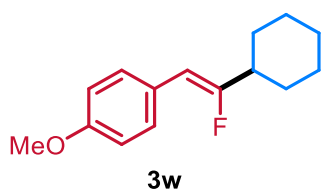
(Z)-1-(2-Fluoro-3-methyl-5-phenylpent-1-en-1-yl)-4-methoxybenzene

Compound **3u** was prepared following the general procedure, purification by column chromatography on silica gel (PE) yielded **3u** (62.2 mg, 73%, *Z/E* > 20:1) as light white solid. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.44 – 7.42 (m, 2H), 7.30 – 7.25 (m, 2H), 7.20 – 7.18 (m, 3H), 6.87 – 6.85 (m, 2H), 5.42 (d, *J* = 40.4 Hz, 1H), 3.81 (s, 3H), 2.73 – 2.60 (m, 2H), 2.44 – 2.31 (m, 1H), 1.99 – 1.89 (m, 1H), 1.77 – 1.68 (m, 1H), 1.20 (d, *J* = 6.8 Hz, 3H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 162.7 (d, *J* = 267.3 Hz), 158.4 (d, *J* = 3.0 Hz), 142.2, 129.7 (d, *J* = 7.5 Hz), 128.5 (d, *J* = 9.0 Hz), 126.7, 126.0, 114.0, 104.8 (d, *J* = 9.0 Hz), 55.4, 37.4 (d, *J* = 25.7 Hz), 35.6, 33.6, 18.2. ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –113.27. HR-MS (ESI) *m/z* calc. for C₁₉H₂₂FO⁺ [M+H]⁺ : 285.1649, found: 285.1651.



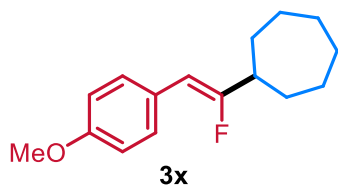
(Z)-1-(2-Cyclopentyl-2-fluorovinyl)-4-methoxybenzene

Compound **3v** was prepared following the general procedure, purification by column chromatography on silica gel (PE) yielded **3v** (33.7 mg, 51%, *Z/E* = 14:1) as colorless liquid. ¹H NMR (600 MHz, Chloroform-*d*) δ 7.43 – 7.41 (m, 2H), 6.86 – 6.85 (m, 2H), 5.44 (d, *J* = 40.2 Hz, 1H), 3.81 (s, 3H), 2.73 – 2.67 (m, 1H), 1.90 – 1.87 (m, 2H), 1.75 – 1.58 (m, 6H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 162.2 (d, *J* = 264.3 Hz), 158.2 (d, *J* = 3.0 Hz), 129.6 (d, *J* = 7.5 Hz), 126.8, 113.9, 103.5 (d, *J* = 9.0 Hz), 55.4, 43.2 (d, *J* = 25.7 Hz), 30.3, 25.6. ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –109.70. HR-MS (ESI) *m/z* calc. for C₁₄H₁₈FO⁺ [M+H]⁺ : 221.1336, found: 221.1325.



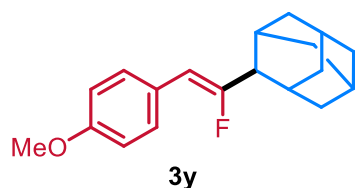
(Z)-1-(2-Cyclohexyl-2-fluorovinyl)-4-methoxybenzene

Compound **3w** was prepared following the general procedure, purification by column chromatography on silica gel (PE) yielded **3w** (46.3 mg, 66%, *Z/E* = 15:1) as colorless liquid. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.42 – 7.26 (m, 2H), 6.86 – 6.84 (m, 2H), 5.37 (d, *J* = 41.1 Hz, 1H), 3.80 (s, 3H), 2.25 – 2.16 (m, 1H), 1.95 – 1.94 (m, 2H), 1.81 – 1.77 (m, 2H), 1.70 – 1.69 (m, 1H), 1.38 – 1.26 (m, 5H). ¹³C NMR (100 MHz, Chloroform-*d*) δ 164.0 (d, *J* = 266.6 Hz), 158.3 (d, *J* = 3.0 Hz), 135.0, 129.7 (d, *J* = 8.0 Hz), 113.9, 103.0 (d, *J* = 9.1 Hz), 55.4, 41.6 (d, *J* = 25.3 Hz), 30.2 (d, *J* = 2.0 Hz), 26.1, 26.1. ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –108.13. HR-MS (ESI) *m/z* calc. for C₁₅H₁₉NaFO⁺ [M+Na]⁺ : 257.1312, found: 257.1322.



(Z)-1-(1-Fluoro-2-(4-methoxyphenyl)vinyl)cycloheptane

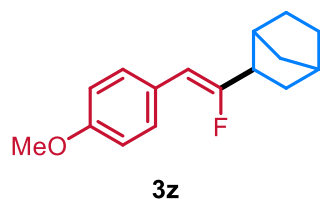
Compound **3x** was prepared following the general procedure, purification by column chromatography on silica gel (PE) yielded **3x** (40.1 mg, 54%, *Z/E* > 20:1) as a colorless liquid. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.41 – 7.39 (m, 2H), 6.86 – 6.84 (m, 2H), 5.40 (d, *J* = 40.8 Hz, 1H), 3.80 (s, 3H), 2.43 – 2.32 (m, 1H), 1.98 – 1.92 (m, 2H), 1.79 – 1.71 (m, 2H), 1.65 – 1.45 (m, 8H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 164.6 (d, *J* = 267.3 Hz), 158.0 (d, *J* = 3.0 Hz), 129.5 (d, *J* = 7.5 Hz), 126.8, 113.8, 102.7 (d, *J* = 9.0 Hz), 55.3, 43.5 (d, *J* = 22.7 Hz), 31.9, 28.3, 26.4. ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –108.28. HR-MS (ESI) *m/z* calc. for C₁₆H₂₂FO⁺ [M+H]⁺ : 249.1649, found: 249.1649.



(Z)-2-(1-Fluoro-2-(4-methoxyphenyl)vinyl)adamantane

Compound **3y** was prepared following the general procedure, purification by column chromatography on silica gel (PE) yielded **3y** (38.6 mg, 45%, *Z/E* > 50:1) as white solid.

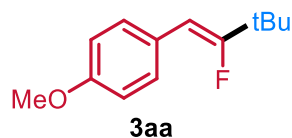
¹H NMR (400 MHz, Chloroform-*d*) δ 7.45 (d, *J* = 8.9 Hz, 2H), 6.86 (d, *J* = 8.8 Hz, 2H), 5.46 (d, *J* = 42.5 Hz, 1H), 3.81 (s, 3H), 2.76 (s, 1H), 2.22 (s, 2H), 2.04 (d, *J* = 12.7 Hz, 2H), 1.93 (d, *J* = 12.6 Hz, 3H), 1.87 – 1.79 (m, 3H), 1.76 (s, 2H), 1.60 (d, *J* = 13.3 Hz, 2H). ¹³C NMR (100 MHz, Chloroform-*d*) δ 162.0 (d, *J* = 266.6 Hz), 158.3 (d, *J* = 3.0 Hz), 129.7 (d, *J* = 8.0 Hz), 114.0, 105.1 (d, *J* = 10.0 Hz), 55.4, 47.0 (d, *J* = 23.2 Hz), 38.6, 37.9, 32.5, 29.4 (d, *J* = 5.0 Hz), 27.9. ¹⁹F NMR (375 MHz, Chloroform-*d*) δ – 108.12. HR-MS (ESI) *m/z* calc. for C₁₉H₂₄FO⁺ [M+H]⁺ : 287.1806, found: 287.1816.



(Z)-2-(1-Fluoro-2-(4-methoxyphenyl)vinyl)bicyclo[2.2.1]heptane

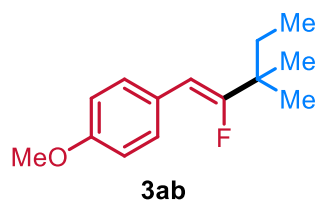
Compound **3z** was prepared following the general procedure, purification by column chromatography on silica gel (PE) yielded **3z** (45.0 mg, 61%, *Z/E* > 20:1) as white solid.

¹H NMR (400 MHz, Chloroform-*d*) δ 7.42 – 7.40 (m, 2H), 6.86 – 6.84 (m, 2H), 5.40 (d, *J* = 40.4 Hz, 1H), 3.80 (s, 3H), 2.40 – 2.29 (m, 2H), 1.62 – 1.50 (m, 1H), 1.57 – 1.52 (m, 4H), 1.30 – 1.17 (m, 4H). ¹³C NMR (100 MHz, Chloroform-*d*) δ 163.0 (d, *J* = 264.6 Hz), 158.3, 129.6 (d, *J* = 7.0 Hz), 113.9, 103.5, 55.4, 45.2 (d, *J* = 25.3 Hz), 40.9 (d, *J* = 2.0 Hz), 36.5 (d, *J* = 2.0 Hz), 36.4, 35.3 (d, *J* = 3.0 Hz), 30.1, 28.8. ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –106.36. HR-MS (ESI) *m/z* calc. for C₁₆H₂₀FO⁺ [M+H]⁺ : 247.1498, found: 247.1502.



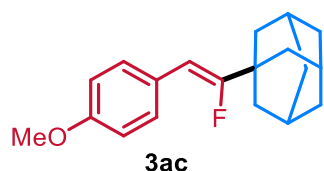
(Z)-1-(2-Fluoro-3,3-dimethylbut-1-en-1-yl)-4-methoxybenzene

Compound **3aa** was prepared following the general procedure, purification by column chromatography on silica gel (PE) yielded **3aa** (53.7 mg, 86%, *Z/E* > 50:1) as colorless liquid. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.45 – 7.44 (m, 2H), 6.87 – 6.85 (m, 2H), 5.47 (d, *J* = 41.1 Hz, 1H), 3.81 (s, 3H), 1.22 (s, 9H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 166.8 (d, *J* = 267.3 Hz), 158.2 (d, *J* = 3.0 Hz), 129.7 (d, *J* = 7.5 Hz), 126.8, 113.9, 101.7 (d, *J* = 10.5 Hz), 55.4, 35.5 (d, *J* = 24.2 Hz), 27.6 (d, *J* = 3.0 Hz). ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –112.36. HR-MS (ESI) *m/z* calc. for C₁₃H₁₇NaFO⁺ [M+Na]⁺ : 231.1156, found: 231.1149.



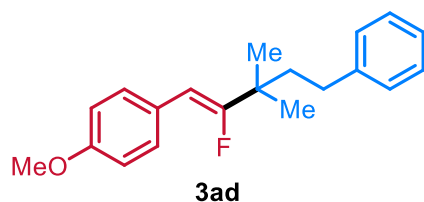
(Z)-1-(2-Fluoro-3,3-dimethylpent-1-en-1-yl)-4-methoxybenzene

Compound **3ab** was prepared following the general procedure, purification by column chromatography on silica gel (PE) yielded **3ab** (45.3 mg, 68%, *Z/E* > 30:1) as colorless liquid. ¹H NMR (600 MHz, Chloroform-*d*) δ 7.43 (d, *J* = 8.4 Hz, 2H), 6.85 (d, *J* = 8.4 Hz, 2H), 5.43 (d, *J* = 41.4 Hz, 1H), 3.80 (s, 3H), 1.54 – 1.51 (m, 2H), 1.15 (s, 6H), 0.87 – 0.86 (m, 3H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 165.6 (d, *J* = 267.3 Hz), 158.3 (d, *J* = 3.0 Hz), 129.8 (d, *J* = 7.5 Hz), 127.0, 113.9, 103.5 (d, *J* = 9.0 Hz), 55.4, 39.0 (d, *J* = 22.7 Hz), 32.5, 25.3 (d, *J* = 4.5 Hz), 9.2. ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –112.80. HR-MS (ESI) *m/z* calc. for C₁₄H₁₉NaFO⁺ [M+Na]⁺ : 245.1312, found: 245.1310.



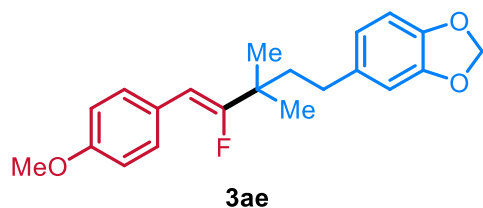
(Z)-1-(1-Fluoro-2-(4-methoxyphenyl)vinyl)adamantane

Compound **3ac** was prepared following the general procedure, purification by column chromatography on silica gel (PE) yielded **3ac** (42.9 mg, 50%, *Z/E* > 50:1) as white solid. ¹H NMR (600 MHz, Chloroform-*d*) δ 7.42 (d, *J* = 8.3 Hz, 2H), 6.85 (d, *J* = 8.3 Hz, 2H), 5.35 (d, *J* = 41.7 Hz, 1H), 3.80 (s, 3H), 2.06 – 2.03 (s, 3H), 1.85 – 1.82 (m, 6H), 1.76 – 1.70 (m, 6H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 167.1 (d, *J* = 265.8 Hz), 158.3 (d, *J* = 3.0 Hz), 129.8 (d, *J* = 7.5 Hz), 127.0, 113.9, 101.8 (d, *J* = 10.6 Hz), 55.4, 39.4 (d, *J* = 1.5 Hz), 36.8, 28.2. ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –117.08. HR-MS (ESI) *m/z* calc. for C₁₉H₂₄FO⁺ [M+H]⁺ : 287.1806, found: 287.1796.



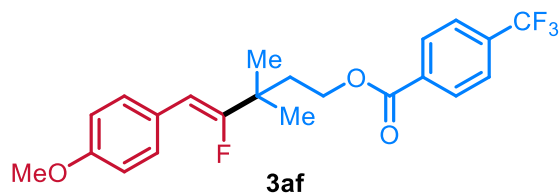
(Z)-1-(2-Fluoro-3,3-dimethyl-5-phenylpent-1-en-1-yl)-4-methoxybenzene

Compound **3ad** was prepared following the general procedure, purification by column chromatography on silica gel (PE) yielded **3ad** (60.8 mg, 68%, *Z/E* > 50:1) as colorless liquid. ¹H NMR (600 MHz, Chloroform-*d*) δ 7.46 – 7.44 (m, 2H), 7.27 – 7.25 (m, 2H), 7.18 – 7.15 (m, 3H), 6.87 – 6.86 (m, 2H), 5.54 (d, *J* = 41.2 Hz, 1H), 3.80 (s, 3H), 2.59 – 2.56 (m, 2H), 1.82 – 1.79 (m, 2H), 1.24 (s, 6H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 165.1 (d, *J* = 267.3 Hz), 158.4, 158.4, 142.8, 129.9 (d, *J* = 7.5 Hz), 128.5, 126.8, 125.8, 114.0, 103.8 (d, *J* = 9.0 Hz), 55.4, 42.2, 38.9 (d, *J* = 22.7 Hz), 31.5, 25.9 (d, *J* = 3.0 Hz). ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –112.94. HR-MS (ESI) *m/z* calc. for C₂₀H₂₄FO⁺ [M+H]⁺ : 299.1806, found: 299.1816.



(Z)-5-(4-Fluoro-5-(4-methoxyphenyl)-3,3-dimethylpent-4-en-1-yl)benzo[d][1,3]dioxole

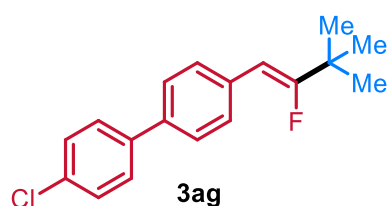
Compound **3ae** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 100:1) yielded **3ae** (74.9 mg, 73%, *Z/E* > 50:1) as colorless liquid. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.47 – 7.45 (m, 2H), 6.98 – 6.87 (m, 2H), 6.73 – 6.68 (m, 2H), 6.6 – 6.62 (m, 1H), 5.91 (s, 2H), 5.51 (d, *J* = 41.4 Hz, 1H), 3.82 (s, 3H), 2.54 – 2.49 (m, 2H), 1.79 – 1.75 (m, 2H), 1.25 (s, 6H). ¹³C NMR (100 MHz, Chloroform-*d*) δ 164.9 (d, *J* = 267.7 Hz), 158.3 (d, *J* = 3.0 Hz), 147.5, 145.5, 136.5, 129.7 (d, *J* = 8.0 Hz), 121.0, 113.8 (d, *J* = 2.0 Hz), 108.9, 108.1, 103.7 (d, *J* = 10.0 Hz), 100.7, 55.3, 42.3, 38.7 (d, *J* = 23.2 Hz), 31.1, 25.7 (d, *J* = 3.0 Hz). ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –112.96. HR-MS (ESI) *m/z* calc. for C₂₁H₂₄FO₃⁺ [M+H]⁺: 343.1704, found: 343.1704.



(Z)-4-Fluoro-5-(4-methoxyphenyl)-3,3-dimethylpent-4-en-1-yl 4-(trifluoromethyl)benzoate

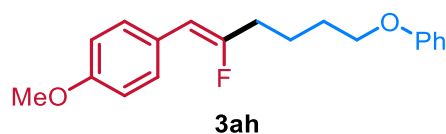
Compound **3af** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 100:1) yielded **3af** (86.1 mg, 70%, *Z/E* > 50:1) as white solid. ¹H NMR (400 MHz, Chloroform-*d*) δ 8.04 (d, *J* = 8.1 Hz, 2H), 7.53 (d, *J* = 8.2 Hz, 2H), 7.35 (d, *J* = 8.8 Hz, 2H), 6.80 (d, *J* = 8.8 Hz, 2H), 5.50 (d, *J* = 41.3 Hz, 1H), 4.44 (t, *J* = 6.6 Hz, 2H), 3.79 (s, 3H), 2.04 (t, *J* = 6.6 Hz, 2H), 1.30 (s, 6H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 165.5, 164.4 (d, *J* = 267.3 Hz), 158.5, 133.4, 134.3 (q, *J* = 33.2 Hz), 130.1, 129.9 (d, *J* = 7.5 Hz), 126.4, 125.3 (d, *J* = 3.0 Hz), 123.8 (q, *J* = 273.3 Hz), 113.9, 103.8 (d, *J* = 9.1 Hz), 62.6, 55.3, 38.1, 37.6 (d, *J* = 27.2 Hz), 26.2.

^{19}F NMR (375 MHz, Chloroform-*d*) δ -63.15, -112.60. HR-MS (ESI) m/z calc. for $\text{C}_{22}\text{H}_{23}\text{F}_4\text{O}_3^+$ $[\text{M}+\text{H}]^+$: 411.1583, found: 411.1578.



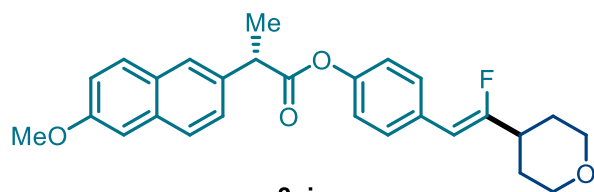
(Z)-4-Chloro-4'-(2-fluoro-3,3-dimethylbut-1-en-1-yl)-1,1'-biphenyl

Compound **3ag** was prepared following the general procedure, purification by column chromatography on silica gel (PE) yielded **3ag** (69.1 mg, 80%, *Z/E* > 50:1) as white solid. ^1H NMR (400 MHz, Chloroform-*d*) δ 7.59 – 7.51 (m, 6H), 7.42 – 7.40 (m, 2H), 5.58 (d, $J = 40.6$ Hz, 1H), 1.26 (s, 9H). ^{13}C NMR (150 MHz, Chloroform-*d*) δ 168.6 (d, $J = 270.3$ Hz), 139.4, 138.1 (d, $J = 3.0$ Hz), 133.7, 133.4, 129.1 (d, $J = 7.5$ Hz), 129.0, 128.3, 127.0, 102.0 (d, $J = 9.0$ Hz), 35.7 (d, $J = 24.2$ Hz), 27.6. ^{19}F NMR (375 MHz, Chloroform-*d*) δ -108.46. HR-MS (ESI) m/z calc. for $\text{C}_{18}\text{H}_{19}\text{ClF}^+$ $[\text{M}+\text{H}]^+$: 289.1159, found: 289.1154.



(Z)-1-(2-fluoro-6-phenoxyhex-1-en-1-yl)-4-methoxybenzene

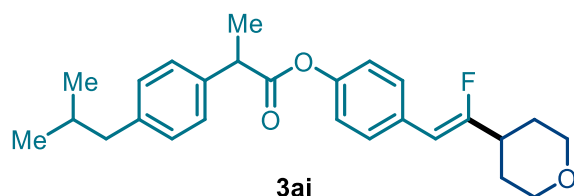
Compound **3ah** was prepared following the general procedure, purification by column chromatography on silica gel (PE) yielded **3ah** (30.6 mg, 34%, *Z/E* > 20:1) as white solid. ^1H NMR (600 MHz, Chloroform-*d*) δ 7.41 – 7.40 (m, 2H), 7.29 – 7.27 (m, 2H), 6.95 – 6.93 (m, 1H), 6.91 – 6.90 (m, 2H), 6.86 – 6.85 (m, 2H), 5.43 (d, $J = 39.8$ Hz, 1H), 4.00 (t, $J = 6.2$ Hz, 2H), 3.81 (s, 3H), 2.42 – 2.37 (m, 2H), 1.89 – 1.86 (m, 2H), 1.81 – 1.78 (m, 2H). ^{13}C NMR (150 MHz, Chloroform-*d*) δ 159.4 (d, $J = 280.9$ Hz), 159.1, 129.7, 129.6, 126.7, 120.8, 114.6, 114.0, 105.6 (d, $J = 9.0$ Hz), 67.5, 55.4, 32.9 (d, $J = 27.2$ Hz), 28.7, 23.2. ^{19}F NMR (375 MHz, Chloroform-*d*) δ -104.21. HR-MS (ESI) m/z calc. for $\text{C}_{19}\text{H}_{22}\text{FO}_2^+$ $[\text{M}+\text{H}]^+$: 301.1598, found: 301.1589.



3ai
from Naproxen

(Z)-4-(2-Fluoro-2-(tetrahydro-2H-pyran-4-yl)vinyl)phenyl (S)-2-(6-methoxy-naphthalen-2-yl)propanoate

Compound **3ai** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 30 :1) yielded **3ai** (87.2 mg, 67%, *Z/E* = 16:1) as white solid. ¹H NMR (600 MHz, Chloroform-*d*) δ 7.77 – 7.73 (m, 3H), 7.51 – 7.49 (m, 1H), 7.44 – 7.43 (m, 2H), 7.18 – 7.14 (m, 2H), 6.95 – 6.94 (m, 2H), 5.42 (d, *J* = 39.9 Hz, 1H), 4.11 – 4.07 (m, 1H), 4.05 – 4.02 (m, 2H), 3.93 (s, 3H), 3.46 – 3.42 (m, 2H), 2.51 – 2.45 (m, 1H), 1.82 – 1.79 (m, 2H), 1.74 – 1.67 (m, 5H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 173.3, 163.2 (d, *J* = 267.3 Hz), 157.9, 149.6 (d, *J* = 3.0 Hz), 135.3, 134.0, 131.4, 129.5 (d, *J* = 7.5 Hz), 129.1, 127.5, 126.3 (d, *J* = 4.5 Hz), 121.5, 119.3, 105.7, 103.7 (d, *J* = 7.5 Hz), 67.6, 55.5, 45.7, 38.8 (d, *J* = 25.7 Hz), 29.8 (d, *J* = 3.0 Hz), 18.7. ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –107.28. HR-MS (ESI) *m/z* calc. for C₂₇H₂₈FO₄⁺ [M+H]⁺ : 435.1966, found: 435.1955.

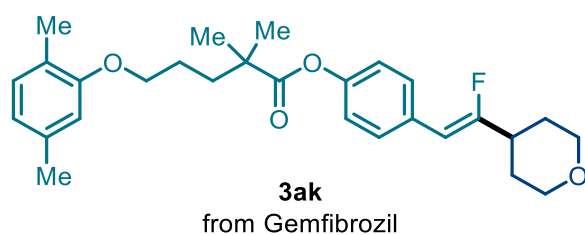


3aj
from Ibuprofen

(Z)-4-(2-Fluoro-2-(tetrahydro-2H-pyran-4-yl)vinyl)phenyl 2-(4-isobutylphenyl)propanoate

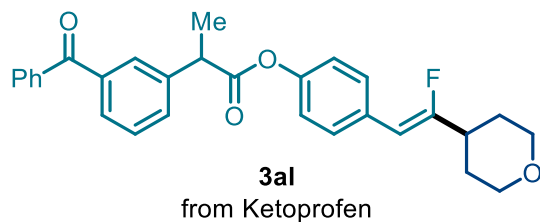
Compound **3aj** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 30:1) yielded **3aj** (73.8 mg, 60%, *Z/E* > 20:1) as white solid. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.45 – 7.43 (m, 2H), 7.31 – 7.29 (m, 2H), 7.15 – 7.13 (m, 2H), 6.96 – 6.94 (m, 2H), 5.43 (d, *J* = 40.0 Hz, 1H), 4.06 – 4.03 (m, 2H), 3.95 – 3.90 (m, 1H), 3.48 – 3.41 (t, *J* = 11.7 Hz, 2H), 2.48 – 2.46 (m, 3H),

1.90– 1.80 (m, 3H), 1.76 – 1.65 (m, 2H), 1.61 – 1.59 (m, 3H), 0.91 (d, $J = 6.6$ Hz, 6H). ^{13}C NMR (100 MHz, Chloroform-*d*) δ 173.3, 163.1 (d, $J = 267.7$ Hz), 149.7 (d, $J = 3.0$ Hz), 141.0, 137.4, 131.3 (d, $J = 2.0$ Hz), 129.7, 129.5 (d, $J = 8.0$ Hz), 127.4, 121.5, 103.8 (d, $J = 8.0$ Hz), 67.6, 45.4, 45.2, 38.9 (d, $J = 25.3$ Hz), 30.3, 29.8 (d, $J = 3.0$ Hz), 22.5, 18.7. ^{19}F NMR (375 MHz, Chloroform-*d*) δ –107.33. HR-MS (ESI) m/z calc. for $\text{C}_{26}\text{H}_{32}\text{FO}_3^+ [\text{M}+\text{H}]^+$: 411.2330, found: 411.2333.



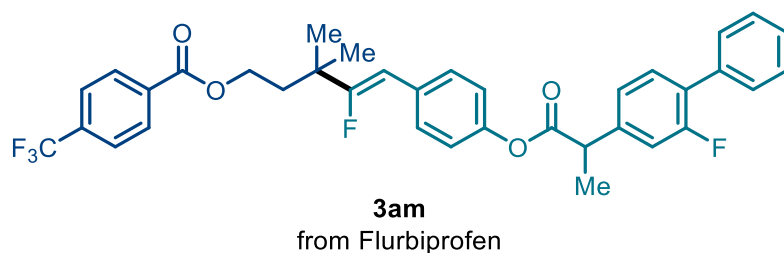
(*Z*)-4-(2-Fluoro-2-(tetrahydro-2*H*-pyran-4-yl)vinyl)phenyl 5-(2,5-dimethylphenoxy)-2,2-dimethylpentanoate

Compound **3ak** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 30:1) yielded **3ak** (70.8 mg, 52%, $Z/E > 20:1$) as white solid. ^1H NMR (600 MHz, Chloroform-*d*) δ 7.47 – 7.46 (m, 2H), 7.01 – 6.97 (m, 3H), 6.67 – 6.66 (m, 1H), 6.62 (s, 1H), 5.45 (d, $J = 40.0$ Hz, 1H), 4.06 – 4.04 (m, 2H), 3.99 – 3.97 (m, 2H), 3.47 – 3.43 (m, 2H), 2.53 – 2.46 (m, 1H), 2.30 (s, 3H), 2.17 (s, 3H), 1.88 – 1.82 (m, 6H), 1.76 – 1.69 (m, 2H), 1.36 (s, 6H). ^{13}C NMR (150 MHz, Chloroform-*d*) δ 176.5, 163.1 (d, $J = 267.3$ Hz), 157.0, 149.8, 136.7, 131.3, 130.5, 129.6 (d, $J = 6.0$ Hz), 123.8, 121.6, 120.9, 112.1, 103.8 ($J = 7.5$ Hz), 67.9, 67.6, 42.6, 38.9 (d, $J = 25.7$ Hz), 37.3, 29.9, 25.4, 25.3, 21.6, 16.0. ^{19}F NMR (375 MHz, Chloroform-*d*) δ –107.35. HR-MS (ESI) m/z calc. for $\text{C}_{28}\text{H}_{36}\text{FO}_4^+ [\text{M}+\text{H}]^+$: 455.2592, found: 455.2599.



(Z)-4-(2-Fluoro-2-(tetrahydro-2H-pyran-4-yl)vinyl)phenyl 2-(3-benzoylphenyl)propanoate

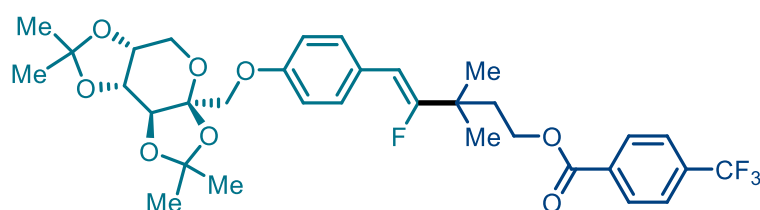
Compound **3al** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 30:1) yielded **3al** (90.7 mg, 66%, *Z/E* > 20:1) as white solid. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.85 – 7.81 (m, 3H), 7.74 – 7.72 (m, 1H), 7.65 – 7.58 (m, 2H), 7.52 – 7.41 (m, 5H), 6.98 – 6.95 (m, 2H), 5.44 (d, *J* = 40.0 Hz, 1H), 4.05 – 4.03 (m, 3H), 3.47 – 3.42 (m, 2H), 2.55 – 2.44 (m, 1H), 1.84 – 1.80 (m, 2H), 1.76 – 1.64 (m, 5H). ¹³C NMR (100 MHz, Chloroform-*d*) δ 196.6, 172.7, 163.3 (d, *J* = 268.7 Hz), 149.4 (d, *J* = 4.0 Hz), 140.5, 138.3, 137.6, 132.7, 131.7, 131.5 (d, *J* = 3.0 Hz), 130.2, 129.6 (d, *J* = 8.0 Hz), 129.4 (d, *J* = 5.1 Hz), 128.9, 128.5, 121.4, 103.7 (d, *J* = 9.0 Hz), 67.6, 45.7, 38.8 (d, *J* = 26.0 Hz), 29.8 (d, *J* = 3.0 Hz), 18.7. ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –107.12. HR-MS (ESI) *m/z* calc. for C₂₉H₂₇NaFO₄⁺ [M+Na]⁺ : 481.1786, found: 481.1774.



(Z)-4-Fluoro-5-(4-((2-(2-fluoro-[1,1'-biphenyl]-4-yl)propanoyl)oxy)phenyl)-3,3-dimethylpent-4-en-1-yl 4-(trifluoromethyl)benzoate

Compound **3am** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 30:1) yielded **3am** (89.6 mg, 48%, *Z/E* > 30:1) as white solid. ¹H NMR (400 MHz, Chloroform-*d*) δ 8.02 – 8.00 (m, 2H), 7.58 – 7.53 (m, 4H), 7.47 – 7.43 (m, 3H), 7.39 – 7.35 (m, 3H), 7.27 – 7.21 (m, 2H), 6.94 – 6.91 (m, 2H), 5.53 (d, *J* = 40.6 Hz, 1H), 4.43 (t, *J* = 6.5 Hz, 2H), 3.99 (q, *J* = 7.1 Hz, 1H), 2.04

(t, $J = 6.5$ Hz, 2H), 1.66 (d, $J = 7.2$ Hz, 3H), 1.30 (s, 6H). ^{13}C NMR (150 MHz, Chloroform-*d*) δ 172.5, 165.8 (d, $J = 270.3$ Hz), 165.5, 160.8, 159.1, 149.4, 141.4 (d, $J = 7.5$ Hz), 135.6, 134.4 (q, $J = 33.2$ Hz), 133.3, 131.5, 131.1 (d, $J = 4.5$ Hz), 130.0, 129.6 (d, $J = 7.5$ Hz), 129.1 (d, $J = 3.0$ Hz), 128.6, 127.9, 125.4 (q, $J = 4.5$ Hz), 123.8 (q, $J = 273.3$ Hz), 123.7 (d, $J = 3.0$ Hz), 121.3, 115.5 (d, $J = 22.7$ Hz), 103.6 (d, $J = 9.0$ Hz), 62.4, 45.3, 38.1, 37.7 (d, $J = 24.2$ Hz), 26.1, 18.5. ^{19}F NMR (375 MHz, Chloroform-*d*) δ -63.10, -109.85, -117.40. HR-MS (ESI) m/z calc. for $\text{C}_{36}\text{H}_{32}\text{F}_5\text{O}_4^+$ $[\text{M}+\text{H}]^+$: 623.2221, found: 623.2231.

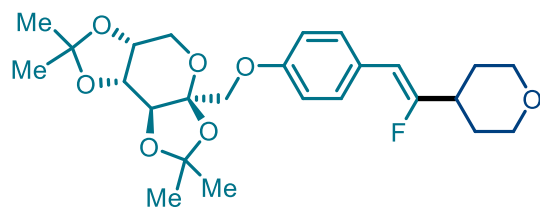


3an

from Diacetonefructose

(*Z*)-4-fluoro-3,3-dimethyl-5-(4-(((3*aS*,5*aR*,8*aR*,8*bS*)-2,2,7,7-tetramethyltetrahydro-3*aH*-bis([1,3]dioxolo)[4,5-*b*:4',5'-*d*]pyran-3*a*-yl)methoxy)phenyl)pent-4-en-1-yl 4-(trifluoromethyl)benzoate

Compound **3an** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 30:1) yielded **3an** (78.5 mg, 41%, *Z/E* > 30:1) as white solid. ^1H NMR (400 MHz, Chloroform-*d*) δ 8.06 – 8.04 (m, 2H), 7.57 – 7.55 (m, 2H), 7.36 – 7.34 (m, 2H), 6.84 – 6.82 (m, 2H), 5.49 (d, $J = 41.3$ Hz, 1H), 4.66 – 4.63 (m, 1H), 4.55 – 4.54 (m, 1H), 4.42 (t, $J = 6.7$ Hz, 2H), 4.28 – 4.26 (m, 1H), 4.16 – 4.13 (m, 1H), 4.04 – 4.02 (m, 1H), 4.00 – 3.96 (m, 1H), 3.81 – 3.78 (m, 1H), 2.03 (t, $J = 6.7$ Hz, 2H), 1.57 (s, 3H), 1.49 – 1.48 (m, 6H), 1.35 (s, 3H), 1.29 (s, 6H). ^{13}C NMR (150 MHz, Chloroform-*d*) δ 165.5, 164.4 (d, $J = 267.3$ Hz), 157.5, 134.4 (q, $J = 31.7$ Hz), 133.5, 130.1, 129.9 (d, $J = 7.5$ Hz), 126.9, 125.4 (d, $J = 3.0$ Hz), 123.8 (q, $J = 271.8$ Hz), 114.7, 109.1 (d, $J = 13.6$ Hz), 103.8 (d, $J = 10.5$ Hz), 102.4, 71.1, 70.4, 70.2, 68.9, 62.6, 61.4, 38.2, 37.6 (d, $J = 24.2$ Hz), 29.9, 26.8, 26.1 (d, $J = 19.6$ Hz), 25.5, 24.2. ^{19}F NMR (375 MHz, Chloroform-*d*) δ -63.10, -112.44. HR-MS (ESI) m/z calc. for $\text{C}_{33}\text{H}_{39}\text{F}_4\text{O}_8^+$ $[\text{M}+\text{H}]^+$: 639.2581, found: 639.2584.

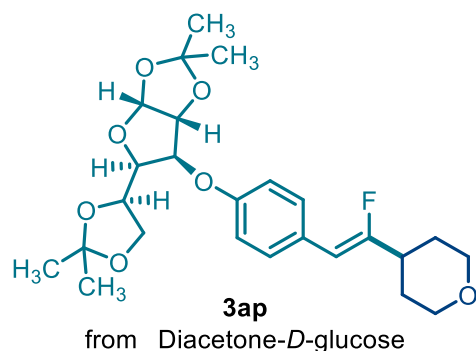


3ao

from Diacetonefructose

(3a*S*,5a*R*,8a*R*,8b*S*)-3a-((4-((*Z*)-2-fluoro-2-(tetrahydro-2*H*-pyran-4-yl)vinyl)phenoxy)methyl)-2,2,7,7-tetramethyltetrahydro-5*H*-bis([1,3]dioxolo)[4,5-*b*:4',5'-*d*]pyran

Compound **3ao** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 30:1) yielded **3ao** (58.5 mg, 42%, *Z/E* = 17:1) as white solid. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.41 – 7.39 (m, 2H), 6.89 – 6.87 (d, *J* = 8.7 Hz, 2H), 5.39 (d, *J* = 40.7 Hz, 1H), 4.65 – 4.63 (m, 1H), 4.55 – 4.53 (m, 1H), 4.28 – 4.26 (m, 1H), 4.17 – 4.14 (m, 1H), 4.05 – 4.02 (m, 3H), 3.99 – 3.95 (m, 1H), 3.81 – 3.77 (m, 1H), 3.48 – 3.41 (m, 2H), 2.53 – 2.43 (m, 1H), 1.83 – 1.80 (m, 2H), 1.76 – 1.67 (m, 2H), 1.57 (s, 3H), 1.48 (s, 6H), 1.34 (s, 3H). ¹³C NMR (100 MHz, Chloroform-*d*) δ 162.0 (d, *J* = 265.6 Hz), 157.5 (d, *J* = 3.0 Hz), 129.8 (d, *J* = 8.0 Hz), 126.8 (d, *J* = 3.0 Hz), 114.7, 109.1 (d, *J* = 8.0 Hz), 103.8 (d, *J* = 9.0 Hz), 102.3, 71.1, 70.3, 70.1, 68.9, 67.6, 61.3, 38.8 (d, *J* = 26.0 Hz), 29.9 (d, *J* = 3.0 Hz), 26.7, 26.1, 25.5, 24.1. ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –109.59. HR-MS (ESI) *m/z* calc. for C₂₅H₃₄FO₇⁺ [M+H]⁺ : 465.2283, found: 465.2288.



(3a*R*,5*R*,6*S*,6a*R*)-5-((*R*)-2,2-dimethyl-1,3-dioxolan-4-yl)-6-(4-((*Z*)-2-fluoro-2-(tetrahydro-2*H*-pyran-4-yl)vinyl)phenoxy)-2,2-dimethyltetrahydrofuro[2,3-*d*][1,3]dioxole

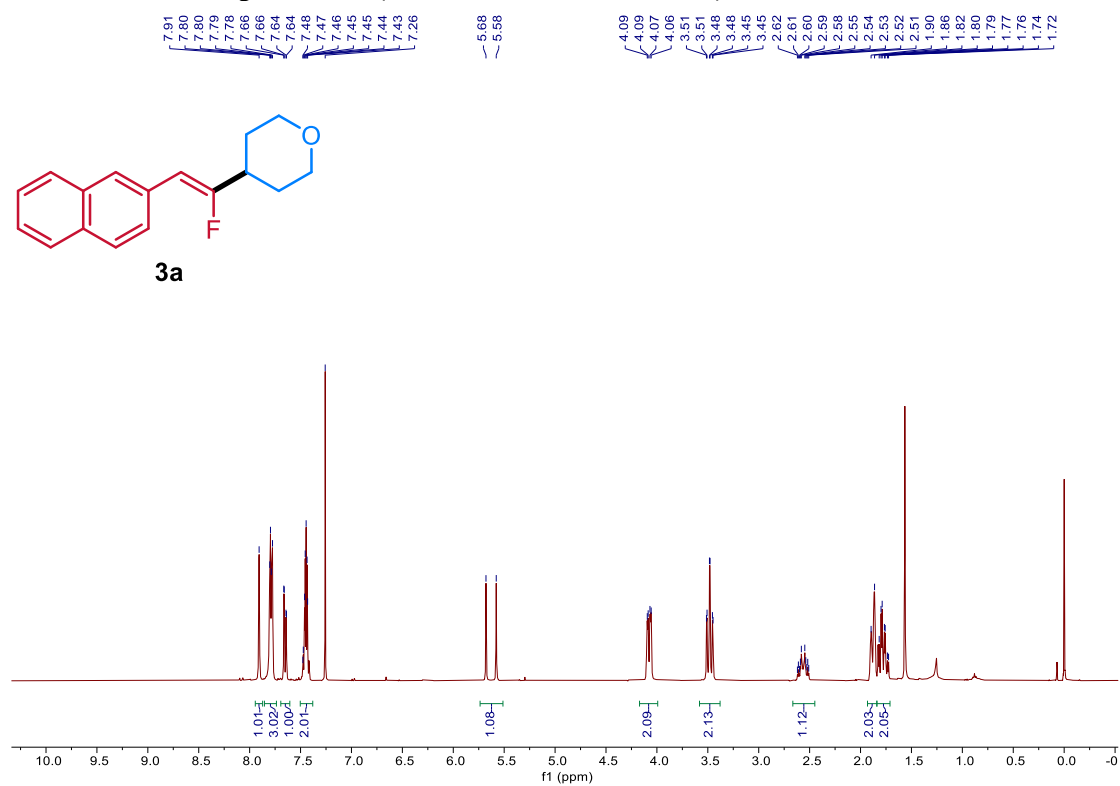
Compound **3ap** was prepared following the general procedure, purification by column chromatography on silica gel (PE/EA = 30:1) yielded **3ap** (69.6 mg, 50%, *Z/E* = 18:1) as white solid. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.44 – 7.42 (m, 2H), 6.95 – 6.92 (m, 2H), 5.93 – 5.92 (m, 1H), 5.41 (d, *J* = 40.4 Hz, 1H), 4.73 – 4.72 (m, 1H), 4.59 – 4.58 (m, 1H), 4.49 – 4.46 (m, 1H), 4.33 – 4.31 (m, 1H), 4.17 – 4.09 (m, 2H), 4.06 – 4.03 (m, 2H), 3.48 – 3.42 (m, 2H), 2.54 – 2.45 (m, 1H), 1.84 – 1.81 (m, 2H), 1.76 – 1.66 (m, 2H), 1.55 (s, 3H), 1.44 (s, 3H), 1.31 (d, *J* = 7.7 Hz, 6H). ¹³C NMR (150 MHz, Chloroform-*d*) δ 162.3 (d, *J* = 264.3 Hz), 155.8 (d, *J* = 1.5 Hz), 130.0 (d, *J* = 7.5 Hz), 127.5, 115.6, 112.2, 109.3, 105.4, 103.7 (d, *J* = 9.0 Hz), 82.3, 80.6, 80.0, 72.4, 67.6, 67.2, 38.8 (d, *J* = 25.6 Hz), 29.9 (d, *J* = 1.5 Hz), 27.0, 26.9, 26.4, 25.4. ¹⁹F NMR (375 MHz, Chloroform-*d*) δ –109.09. HR-MS (ESI) *m/z* calc. for C₂₅H₃₄FO₇⁺ [M+H]⁺ : 465.2283, found: 465.2292.

Reference

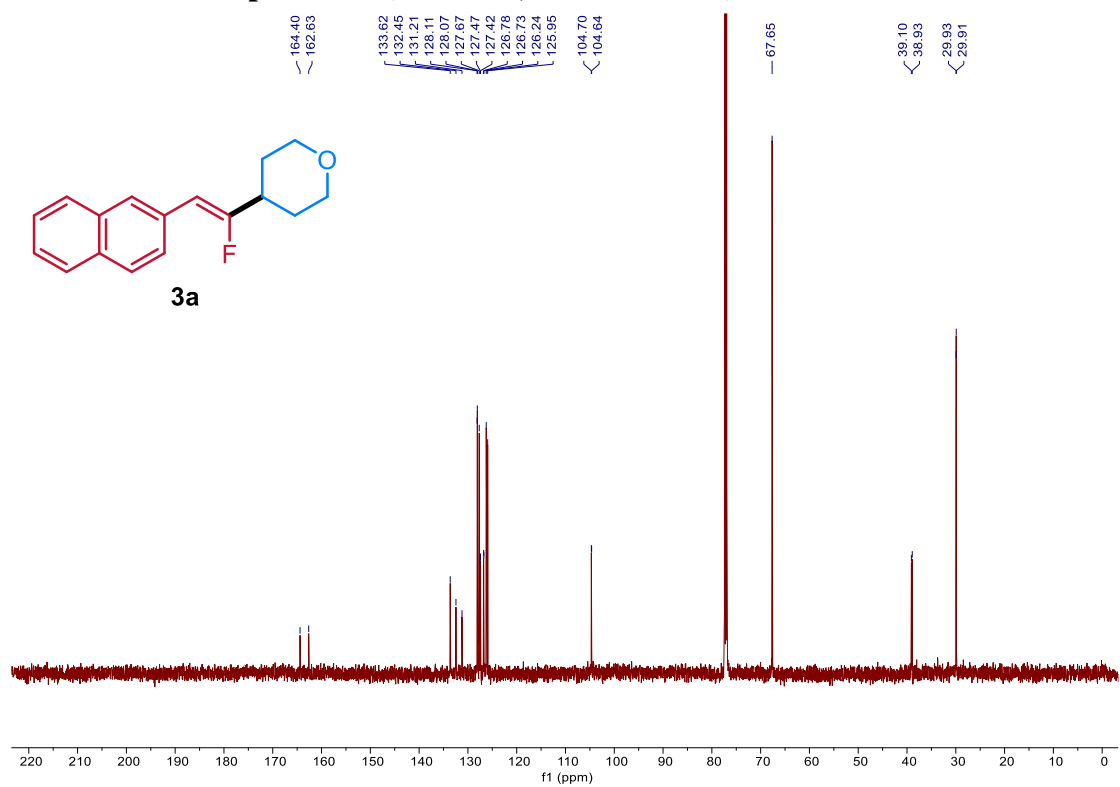
1. S. T. Charles, M. Henry, R. D. Jr. William, *J. Fluorine Chem.*, 2013, **150**, 53–59.
2. J. Yamazaki, T. Watanabe, K. Tanaka, *Tetrahedron Asymmetry.*, 2001, **12**, 669–675.
3. T. Nakamura et al., *Bioorg. Med. Chem. Lett.*, 2004, **14**, 333–336.
4. A. S. Dudnik, G. C. Fu, *J. Am. Chem. Soc.*, 2012, **134**, 10693–10697.
5. B. T. Sargent, E. J. Alexanian, *J. Am. Chem. Soc.*, 2016, **138**, 7520–7523.
6. D. Jan, L. Delaude, F. Simal, A. Demonceau, A. F. Noels. *J. Organomet. Chem.*, 2000, **606**, 55–64.
7. S.-L. Xie, X.-T. Gao, H.-H. Wu, F. Zhou and J. Zhou, *Org. Lett.*, 2020, **22**, 8424–8429.
8. Q. Shao, Y. Huang, *Chem. Commun.*, 2015, **51**, 6584–6586.

NMR Spectra

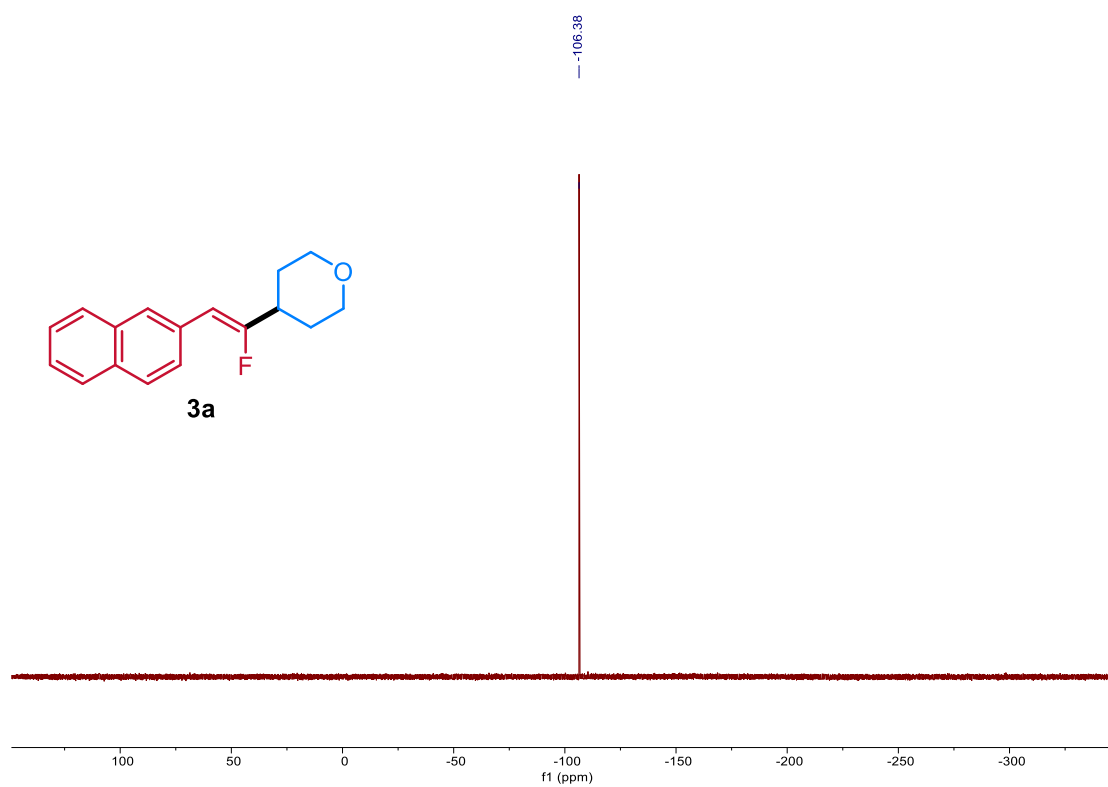
¹H NMR of Compound 3a (400 MHz, Chloroform-*d*):



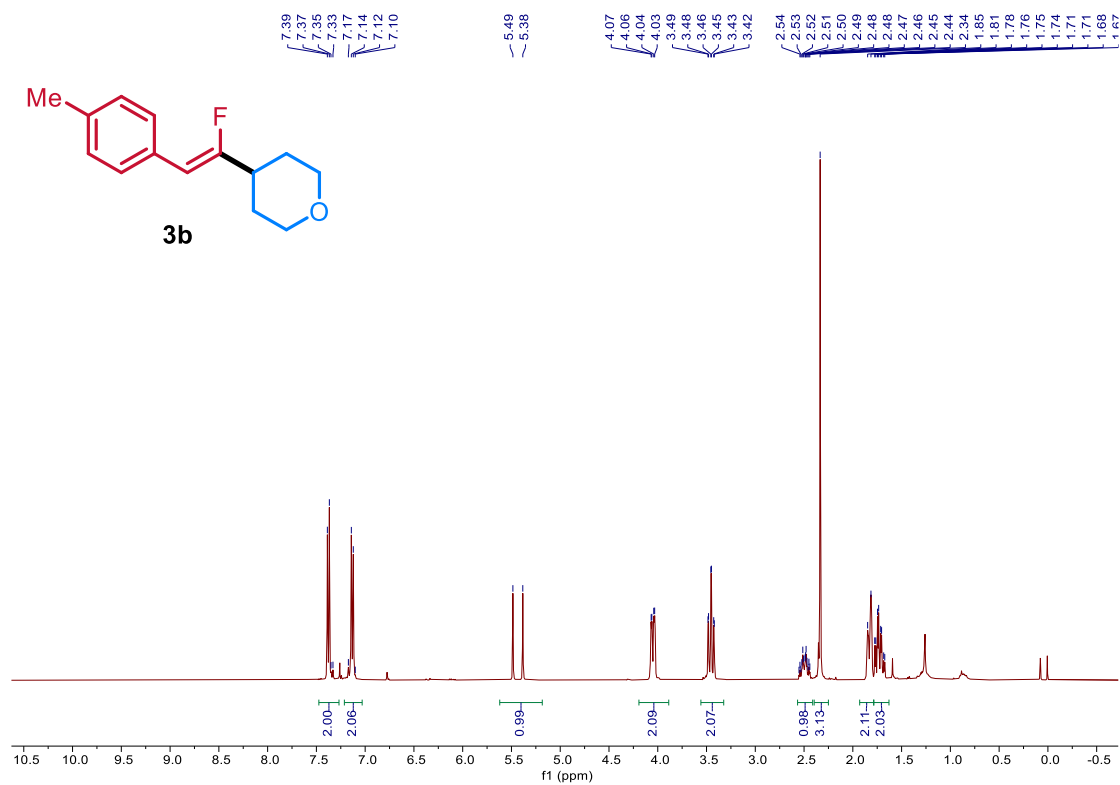
¹³C NMR of Compound 3a (150 MHz, Chloroform-*d*):



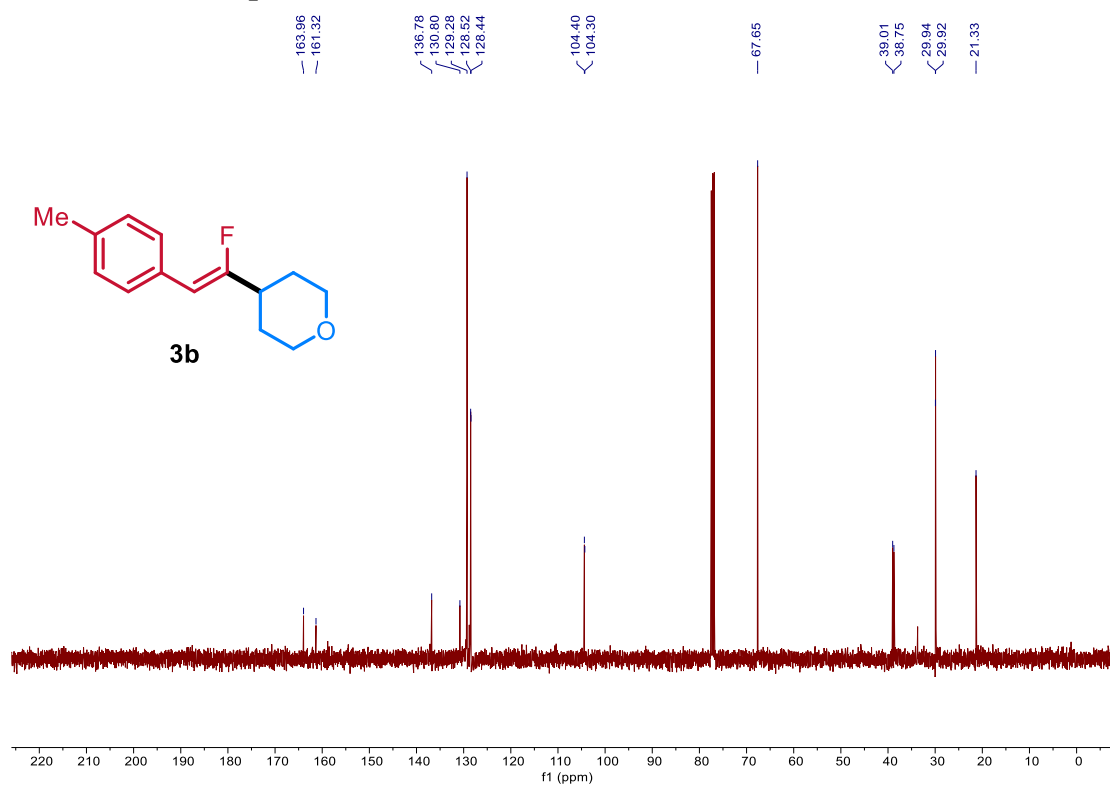
¹⁹F NMR of 3a (375 MHz, Chloroform-*d*):



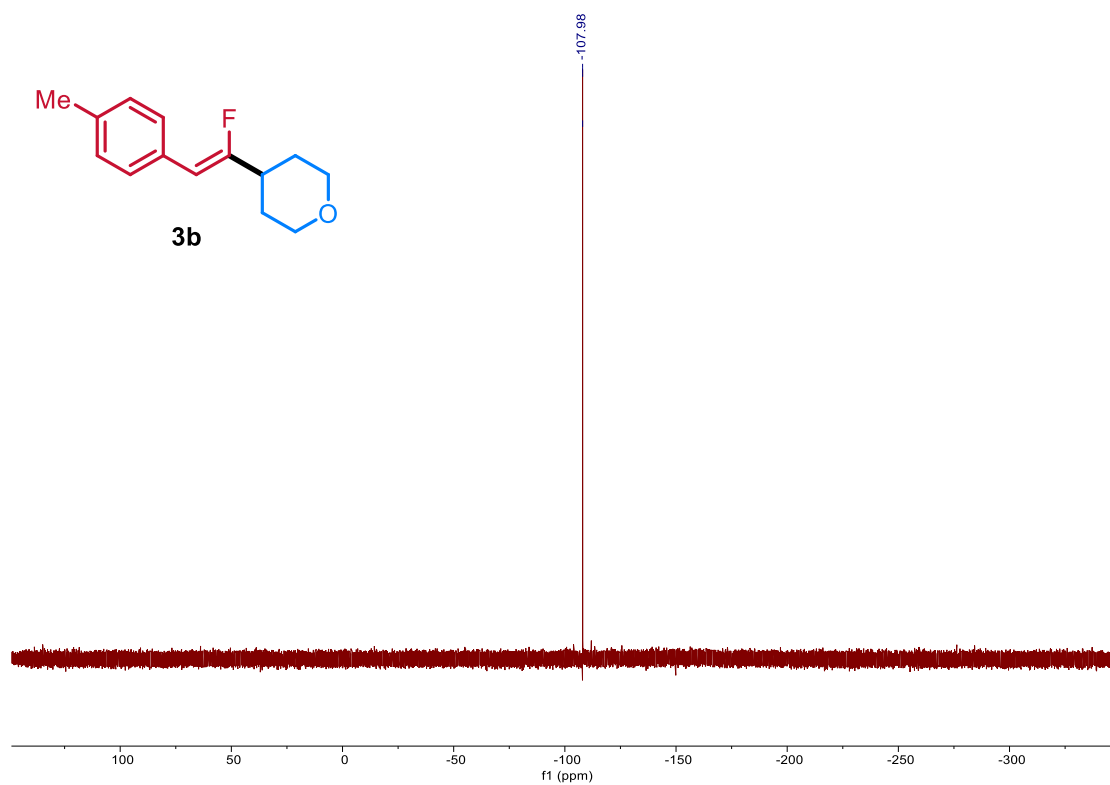
¹H NMR of Compound 3b (400 MHz, Chloroform-*d*):



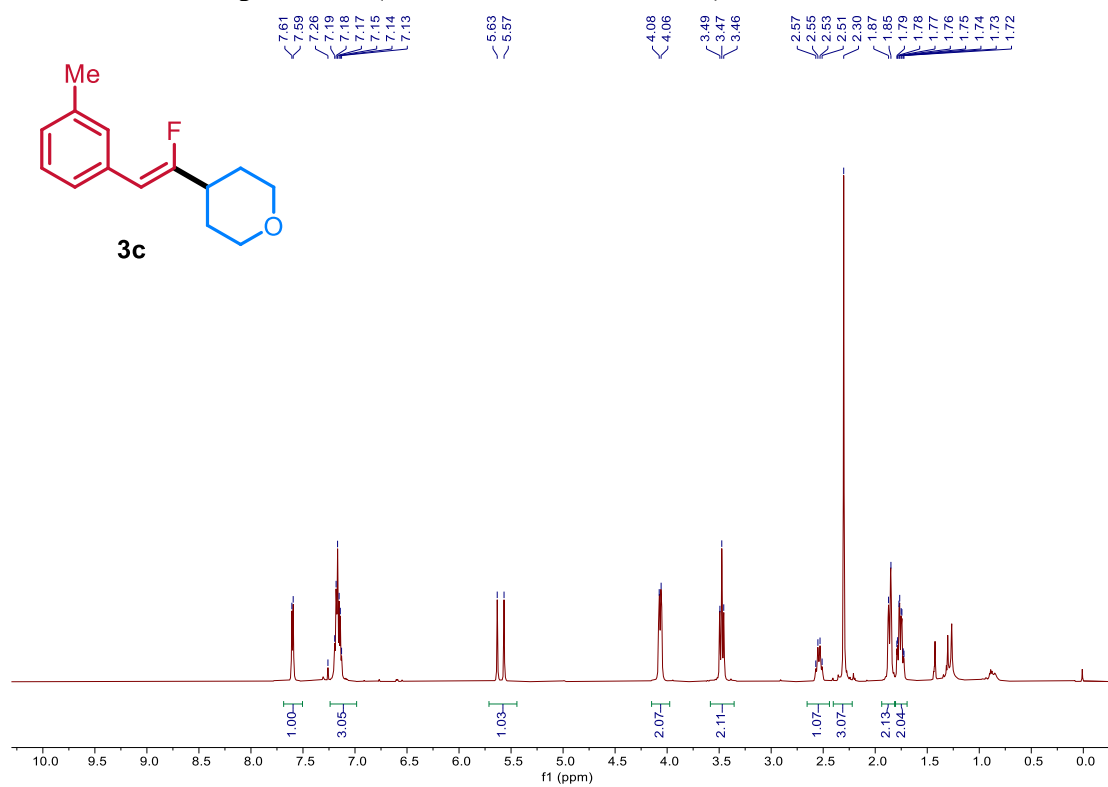
¹³C NMR of Compound 3b (100 MHz, Chloroform-*d*):



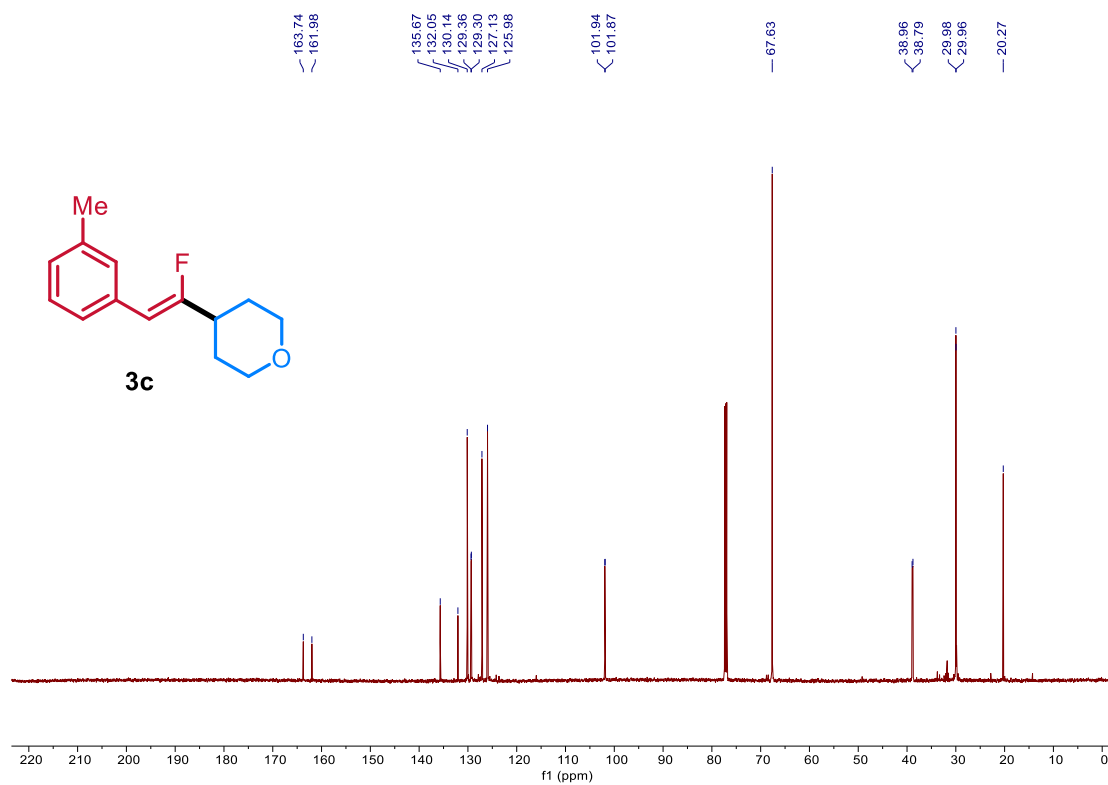
¹⁹F NMR of 3b (375 MHz, Chloroform-*d*):



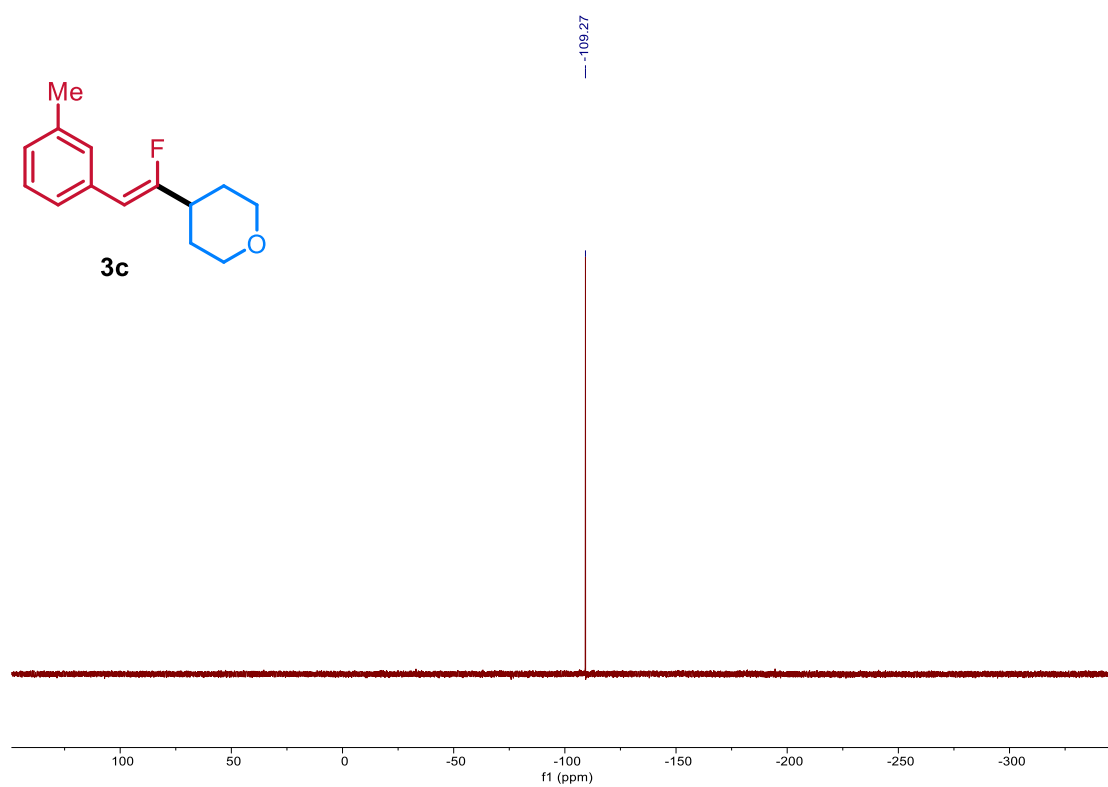
¹H NMR of Compound 3c (400 MHz, Chloroform-*d*):



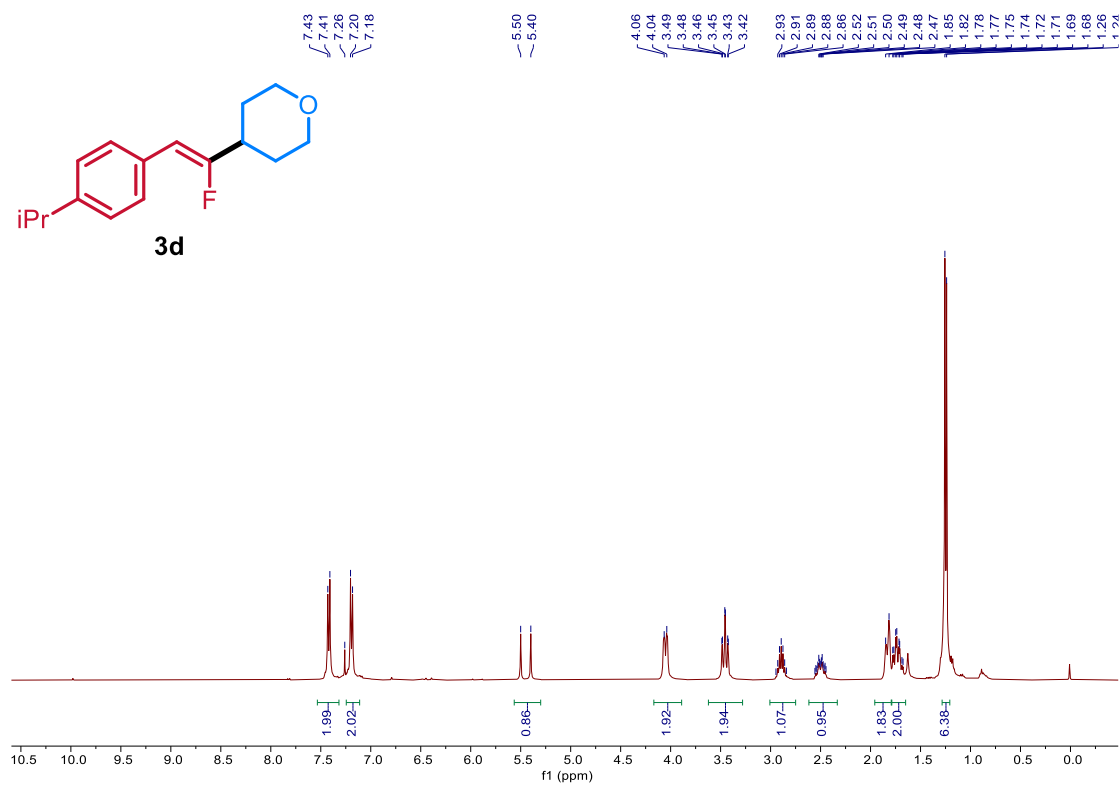
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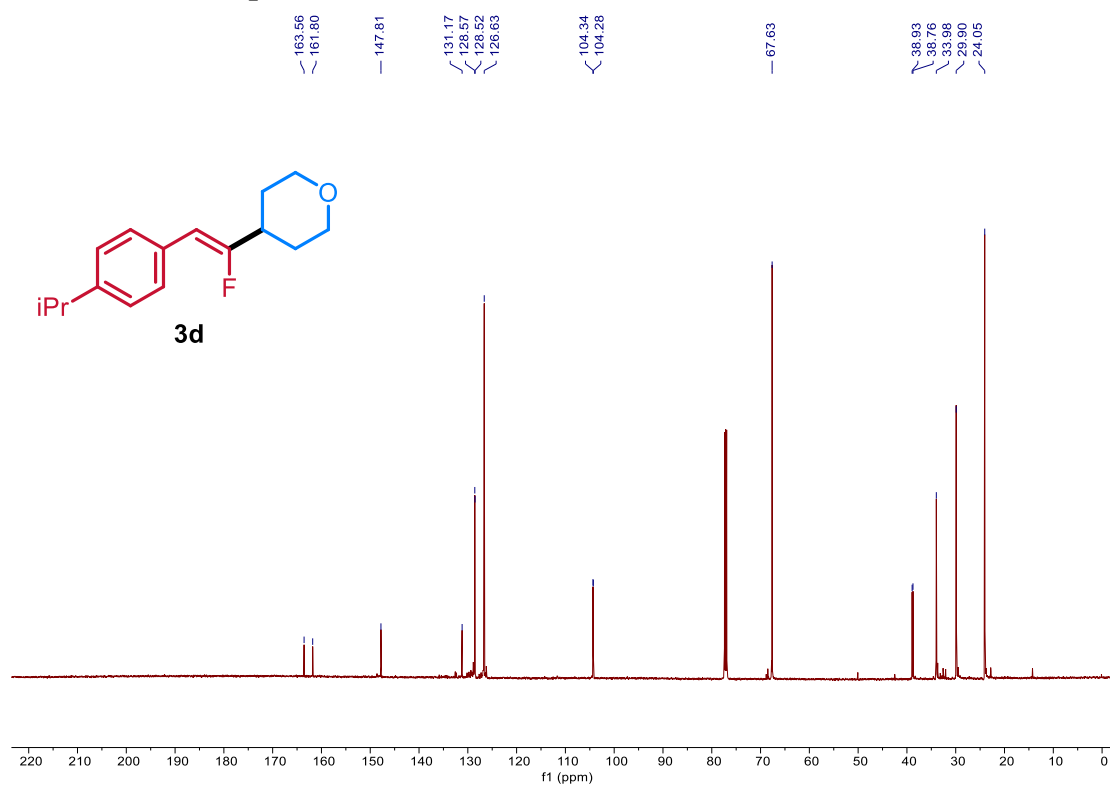
^{19}F NMR of 3c (375 MHz, Chloroform-*d*):



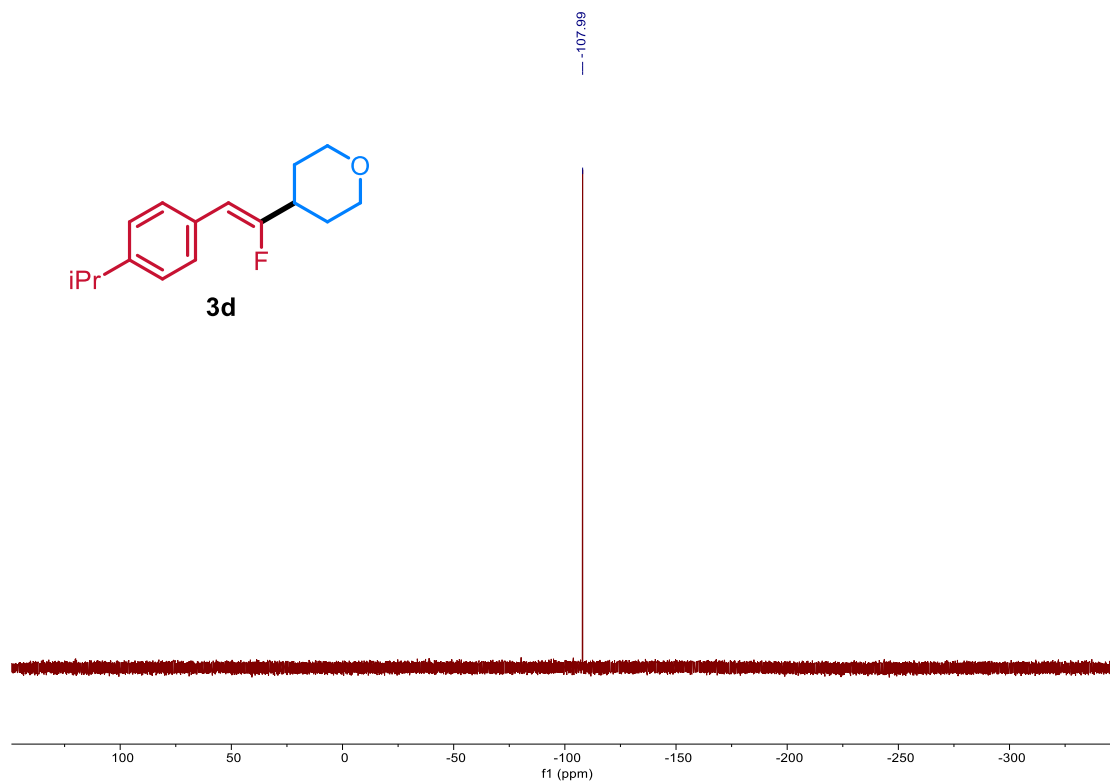
^1H NMR of Compound 3d (400 MHz, Chloroform-*d*):



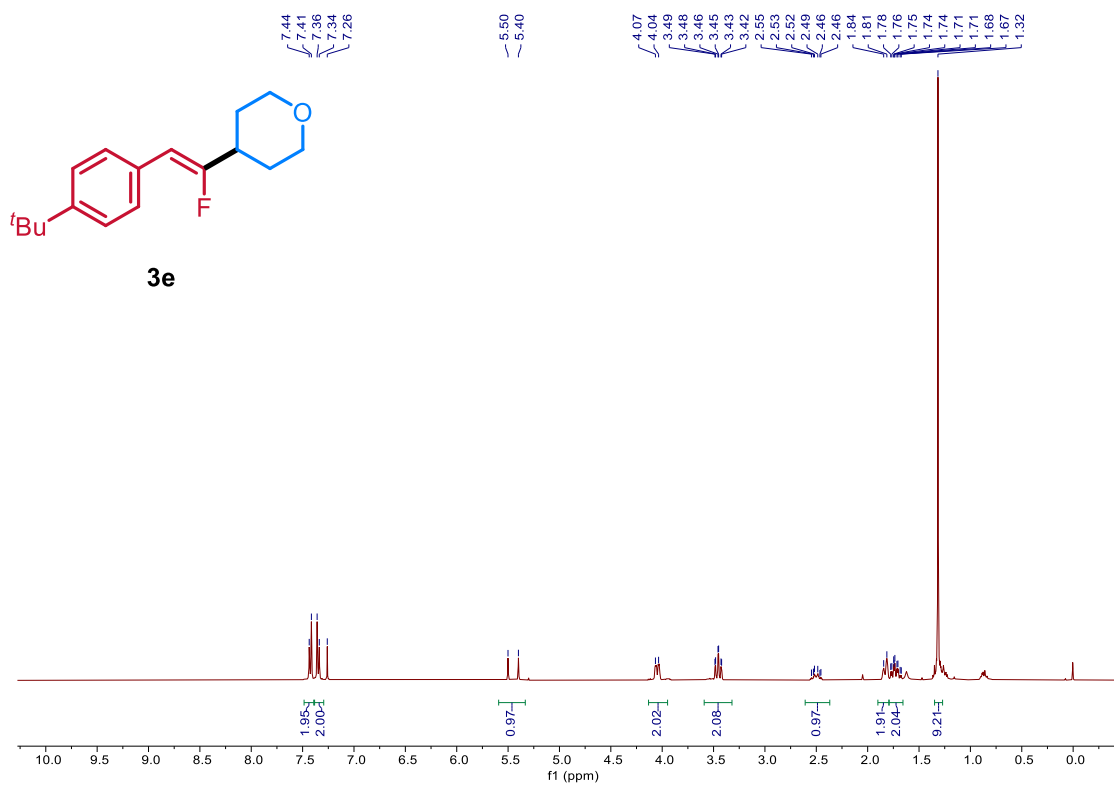
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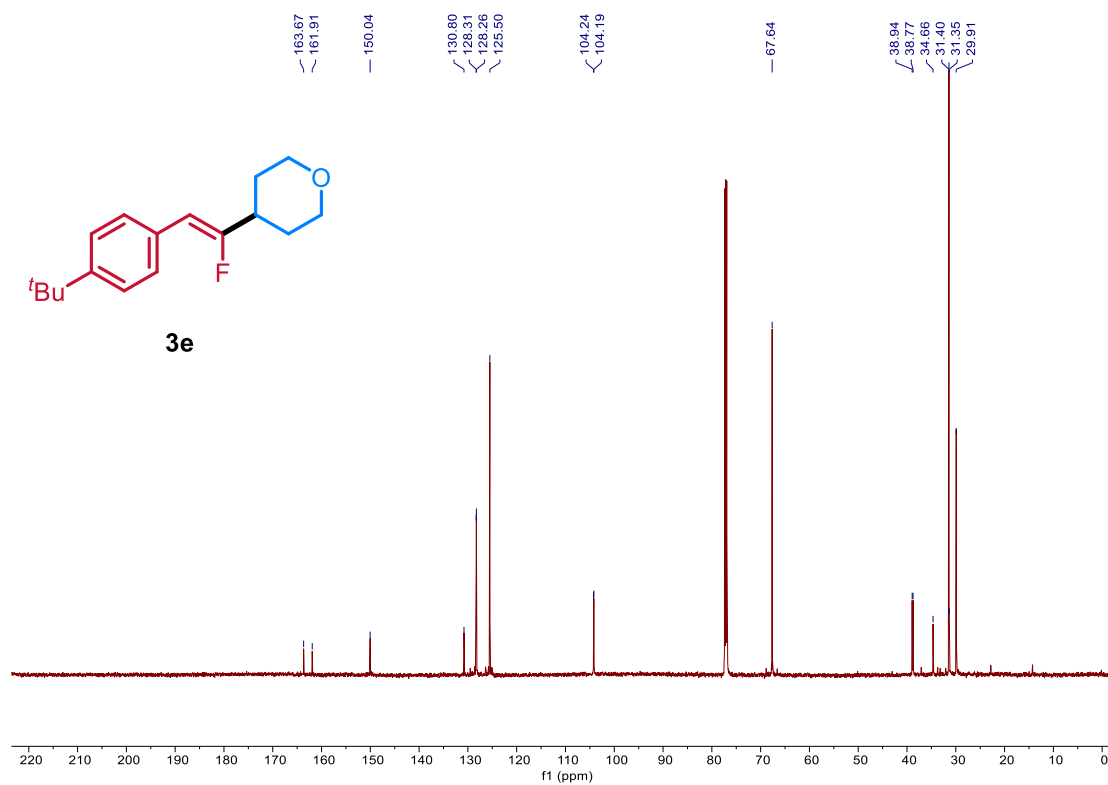
¹⁹F NMR of Compound 3d (375 MHz, Chloroform-*d*):



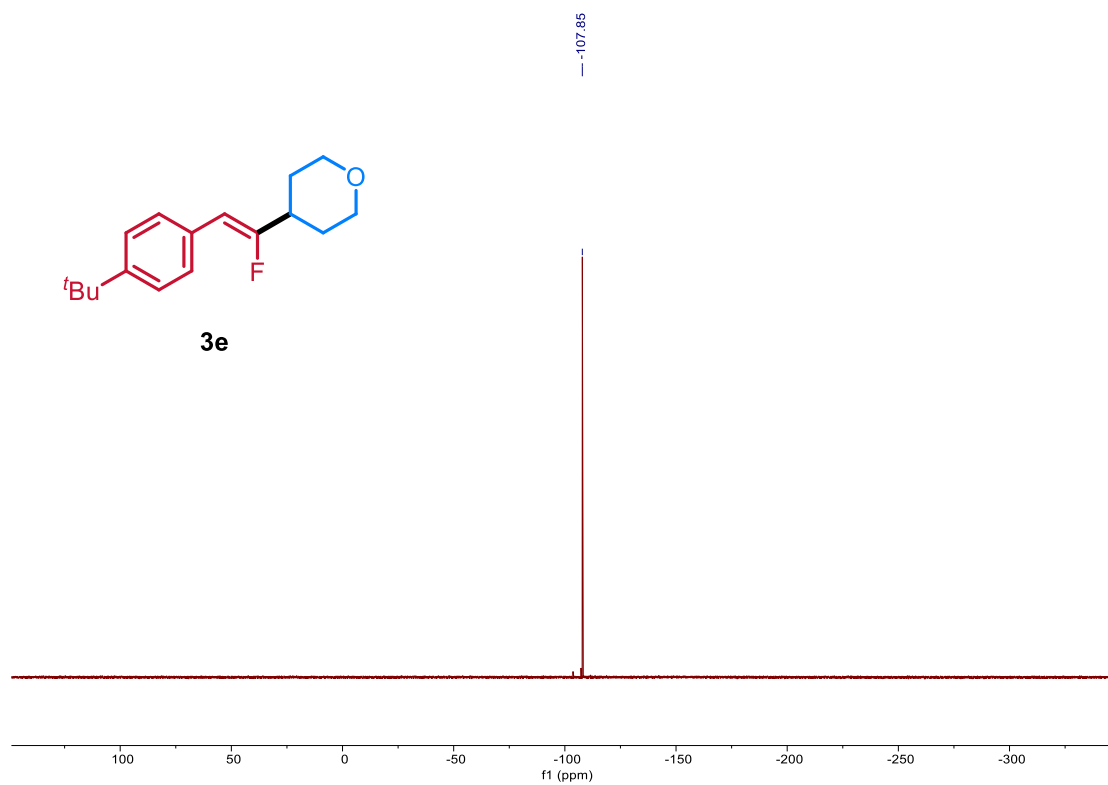
¹H NMR of Compound 3e (400 MHz, Chloroform-*d*):



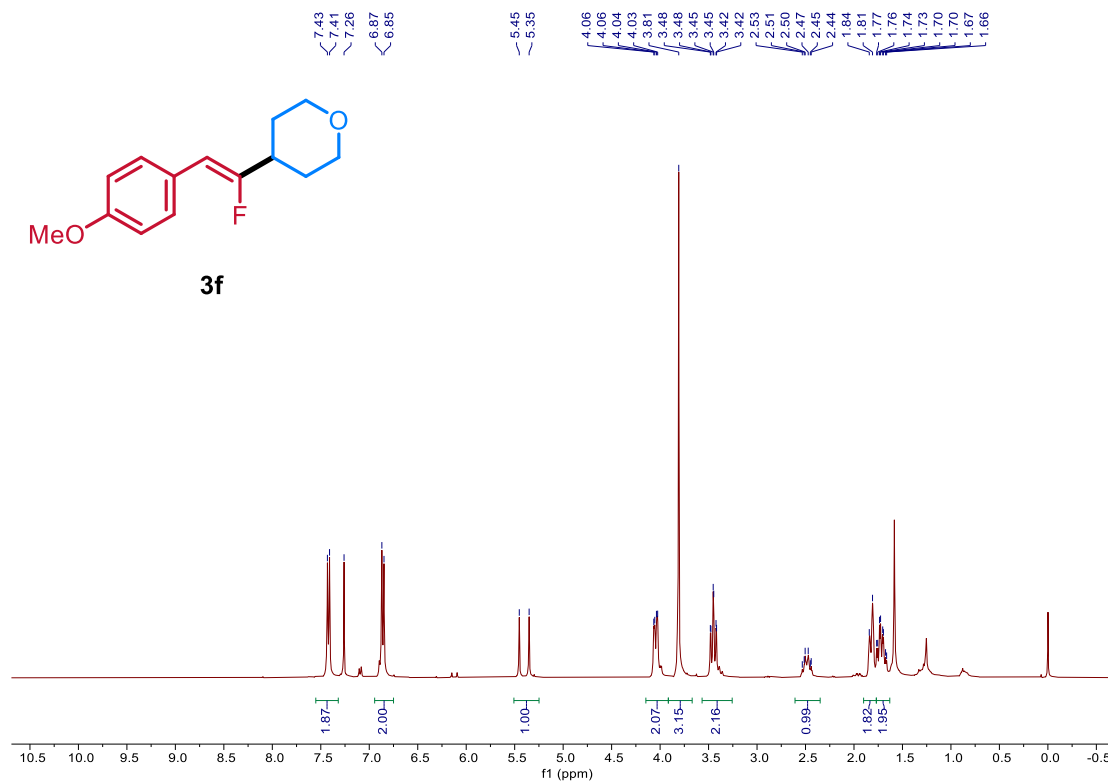
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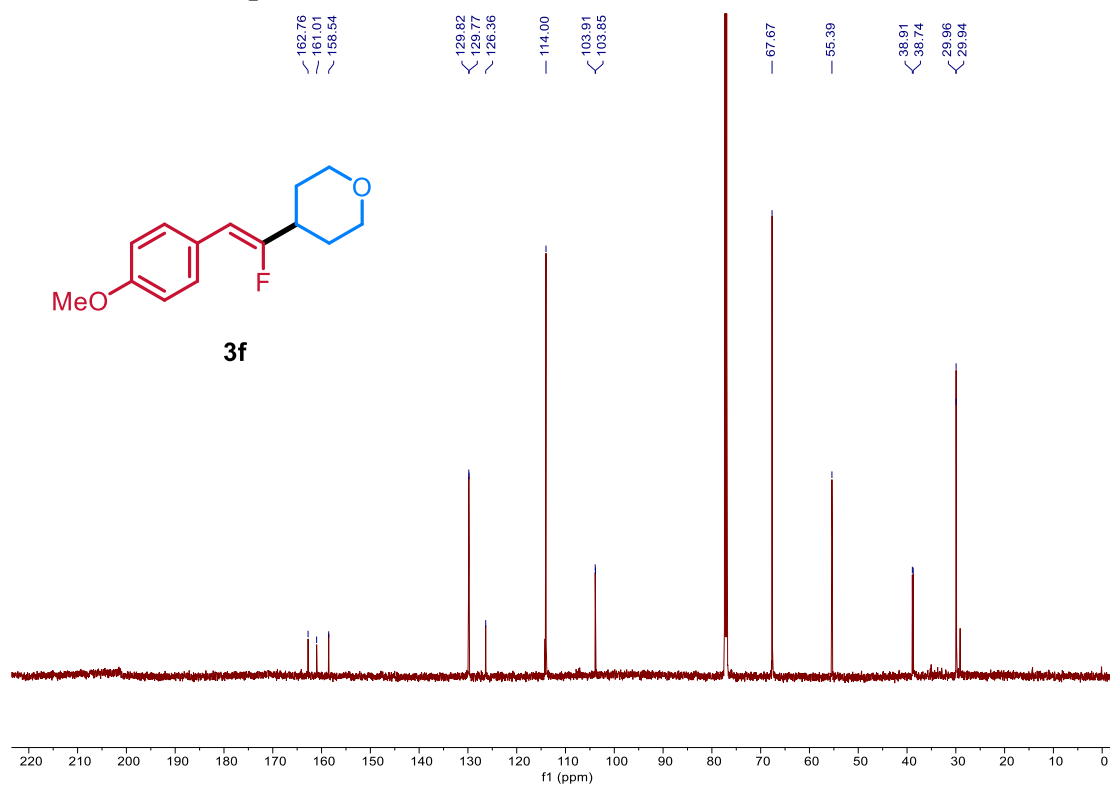
^{19}F NMR of Compound 3e (375 MHz, Chloroform-*d*):



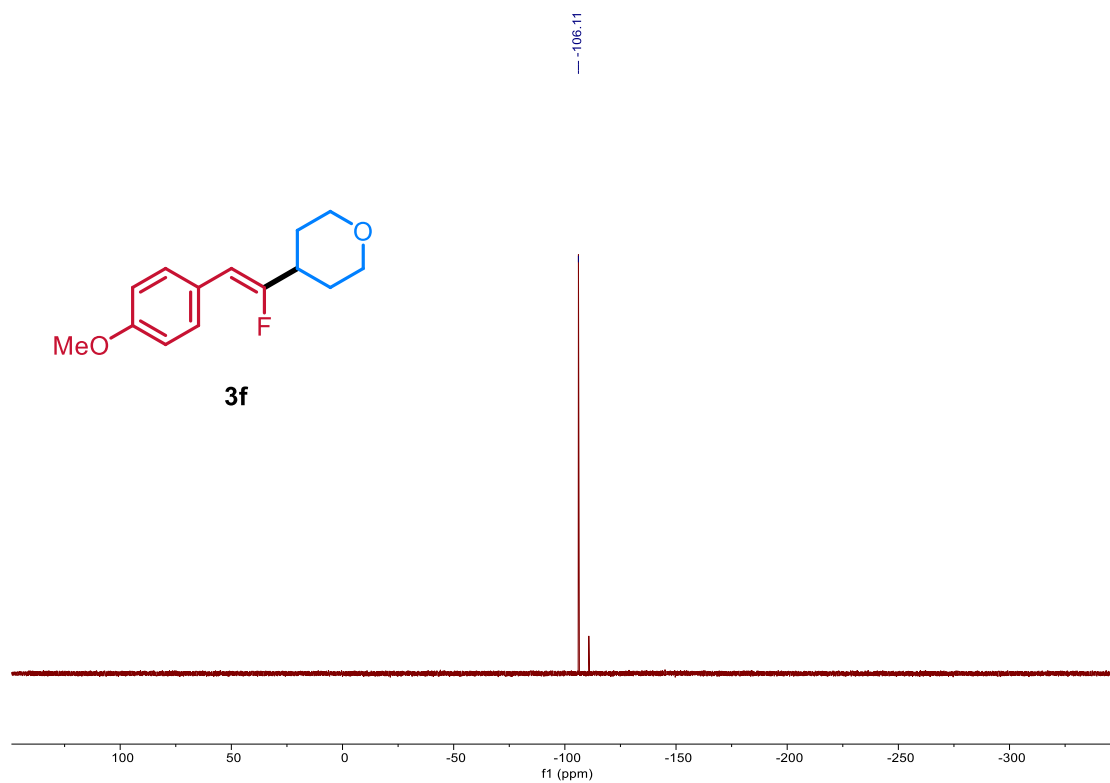
^1H NMR of Compound 3f (400 MHz, Chloroform-*d*):



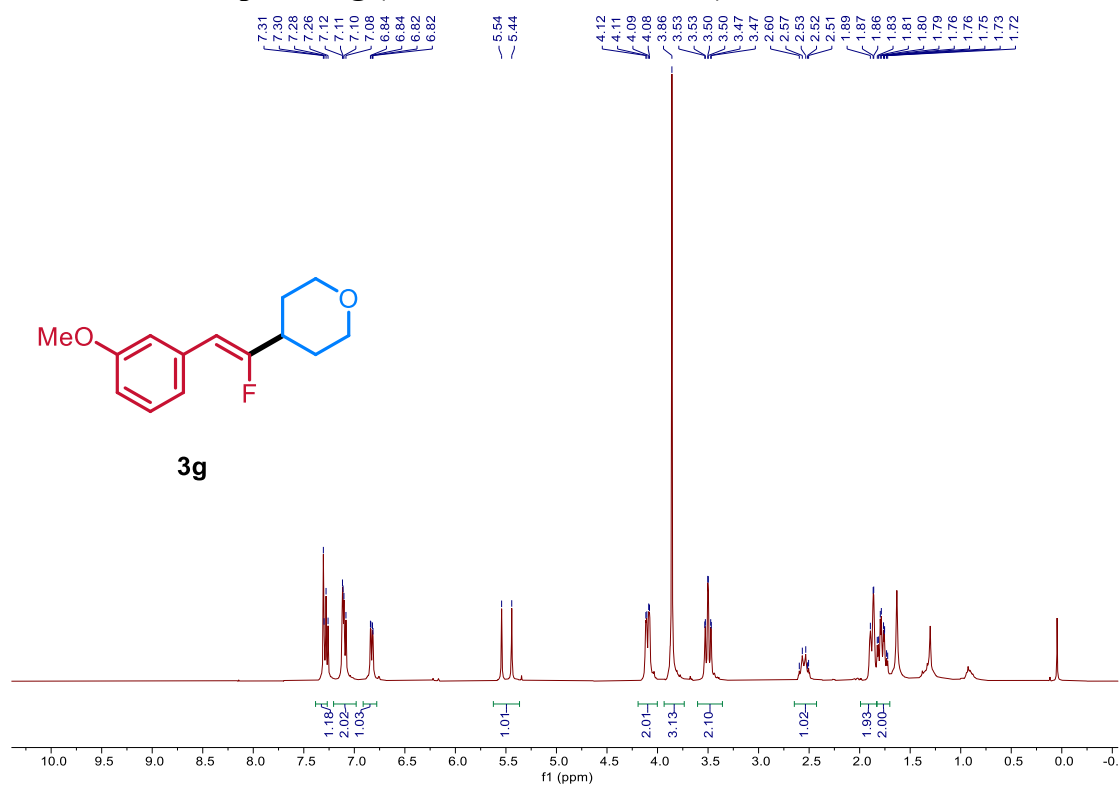
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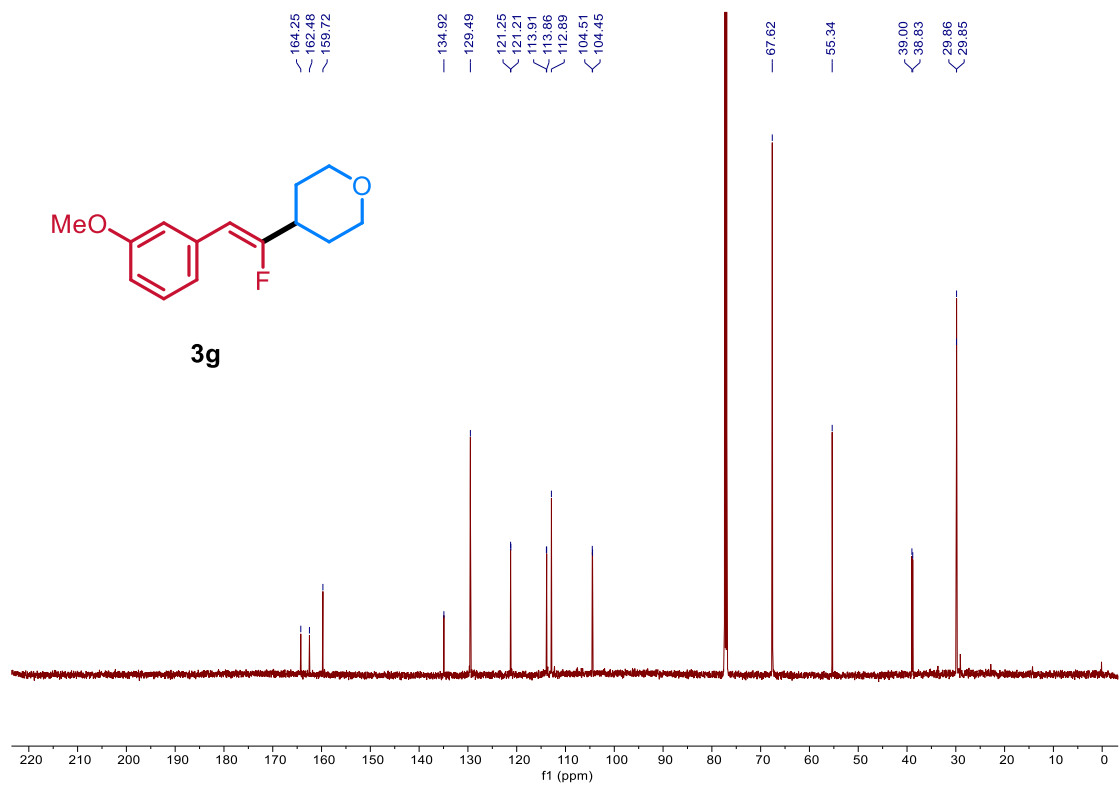
¹⁹F NMR of Compound 3f (375 MHz, Chloroform-*d*):



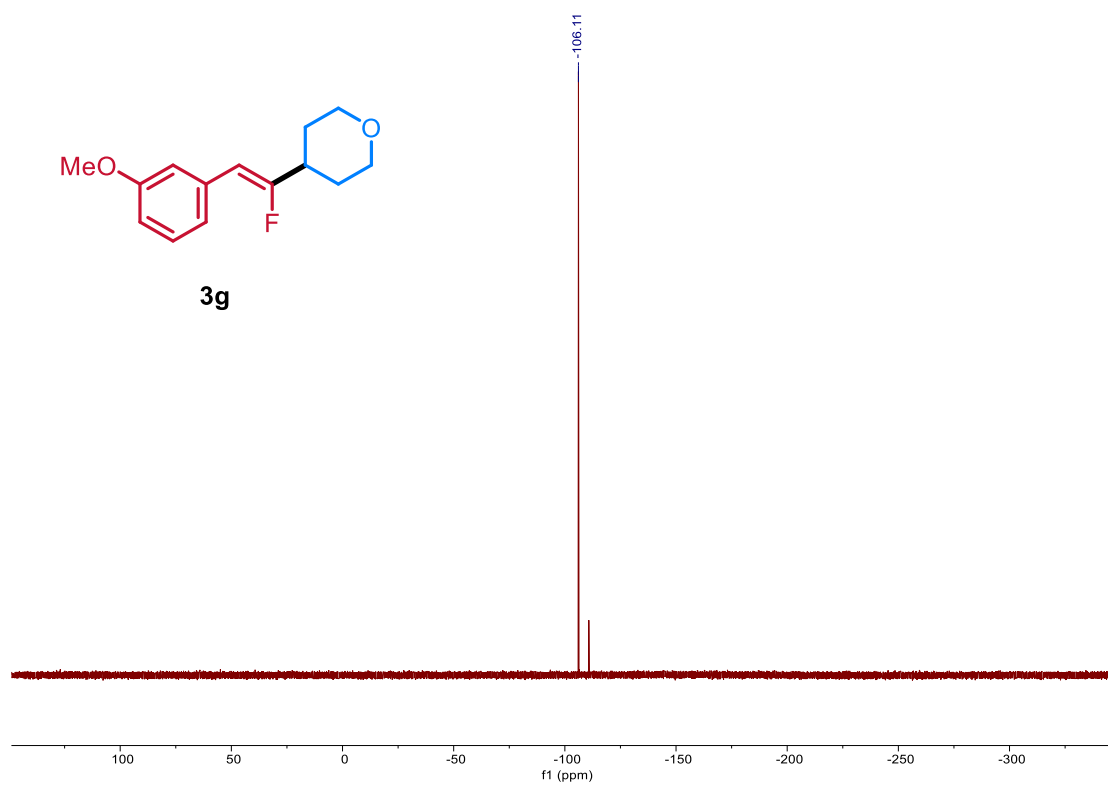
¹H NMR of Compound 3g (400 MHz, Chloroform-*d*):



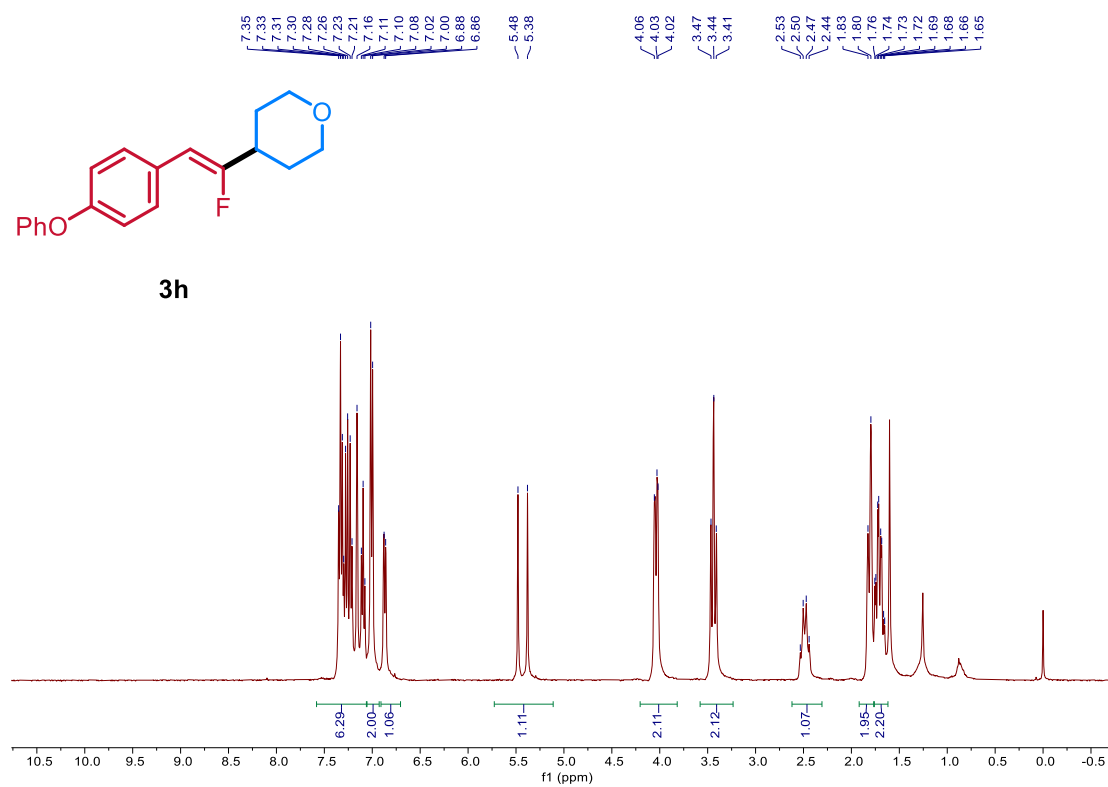
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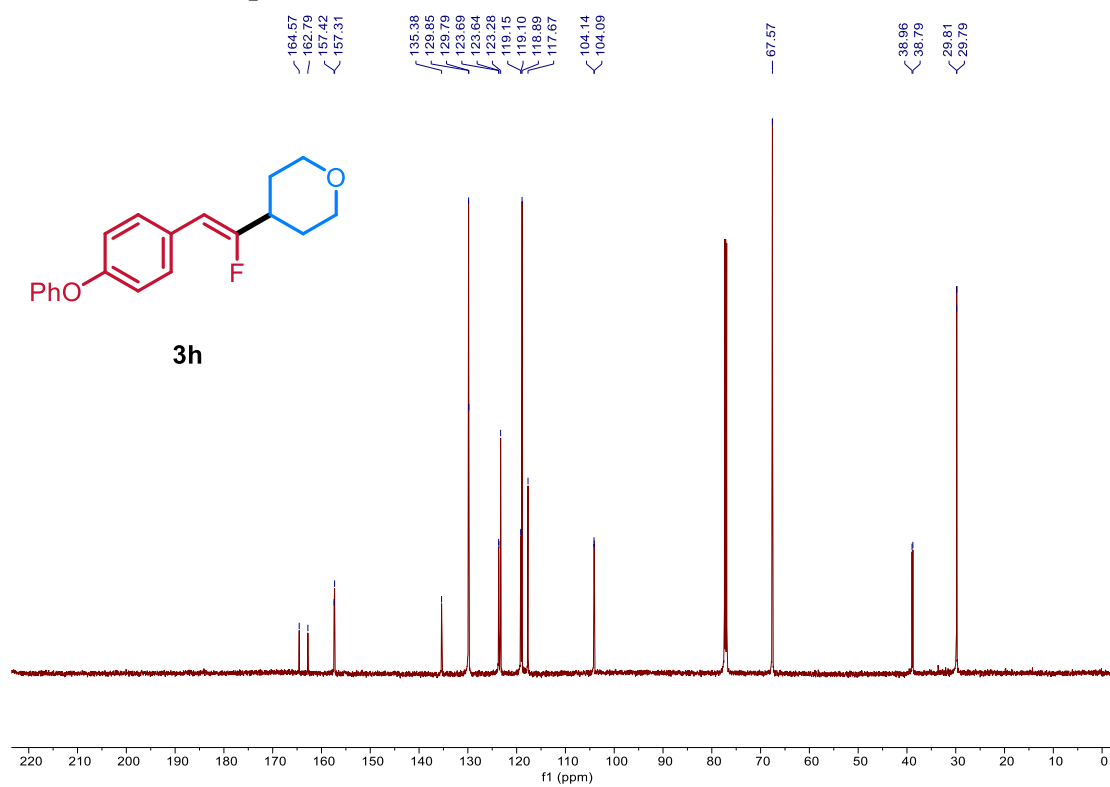
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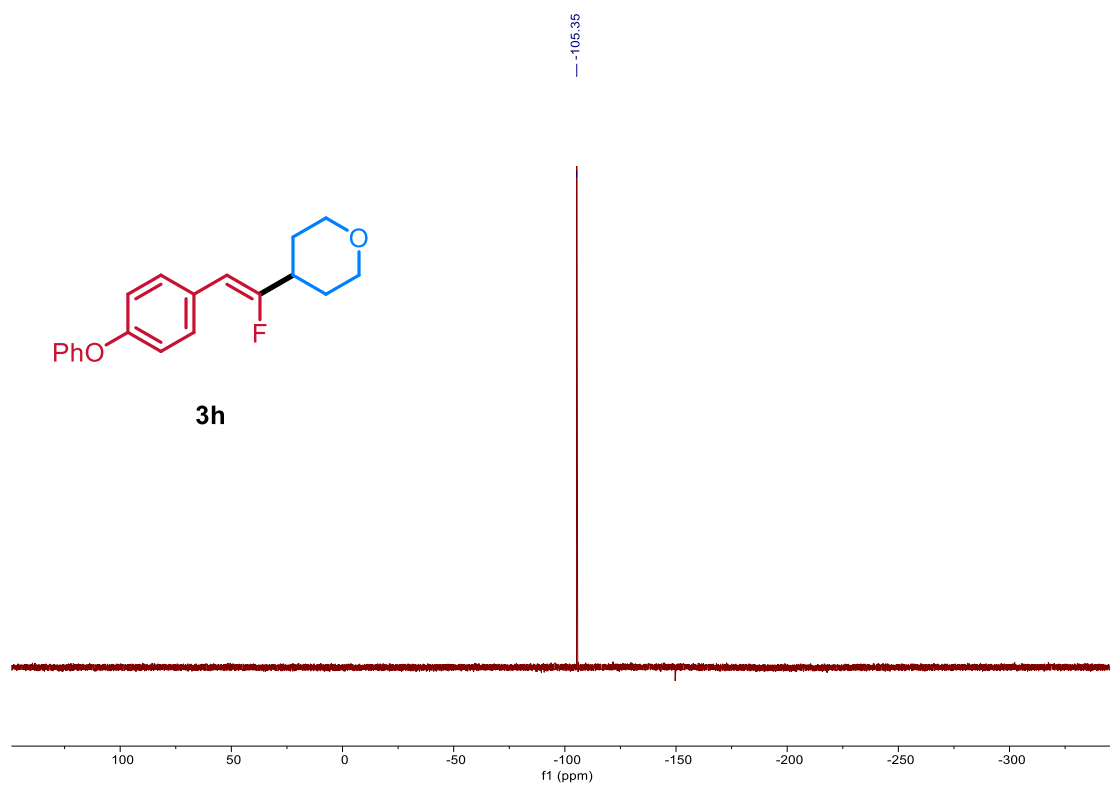
¹H NMR of Compound 3h (400 MHz, Chloroform-*d*):



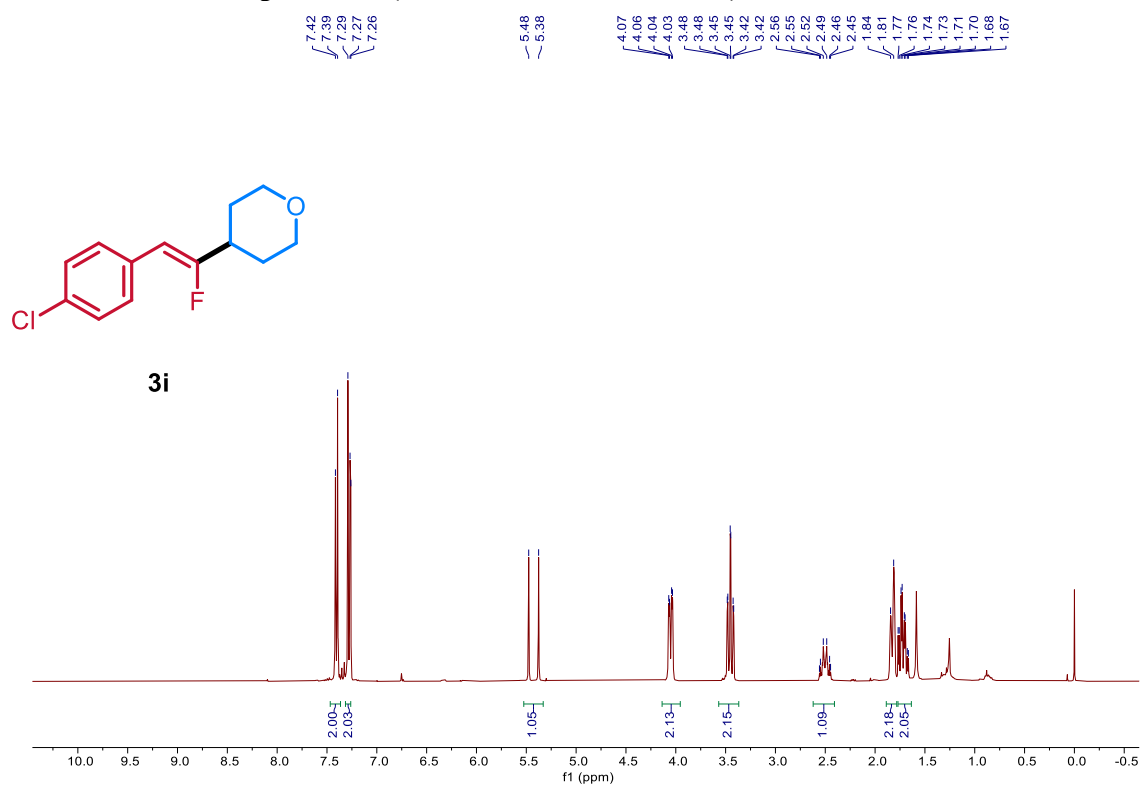
¹³C NMR of Compound 3h (150 MHz, Chloroform-*d*):



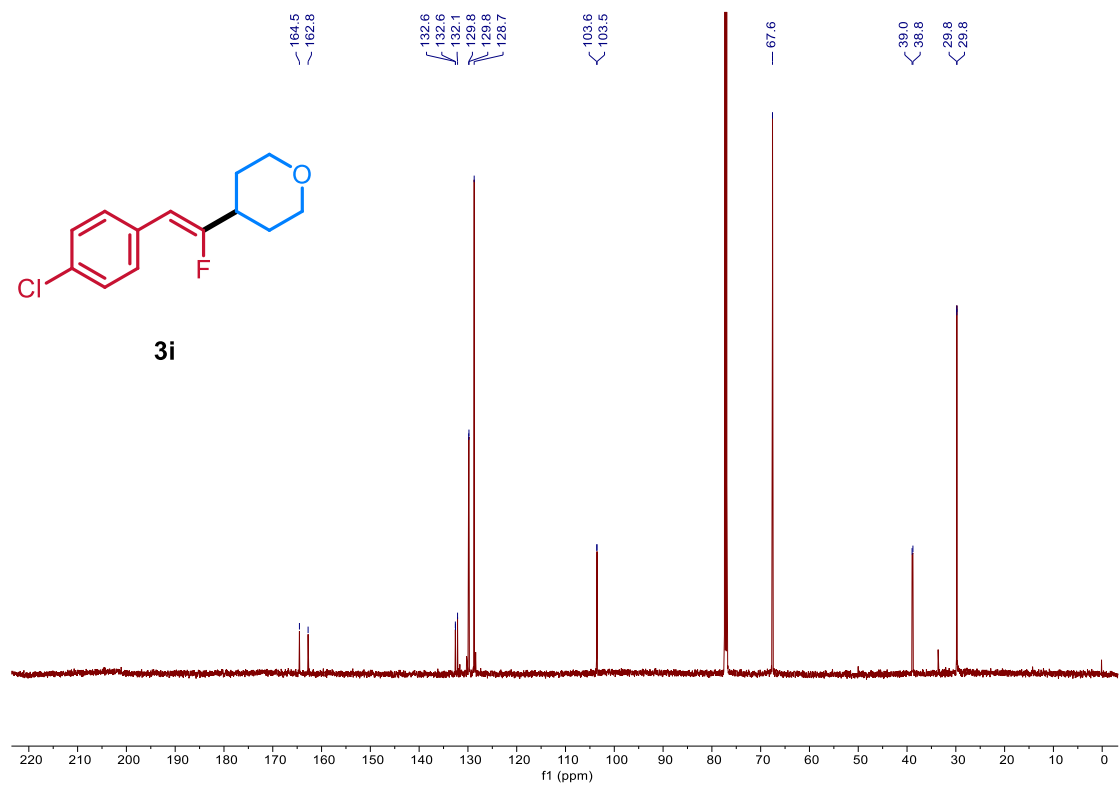
¹⁹F NMR of Compound 3g (375 MHz, Chloroform-*d*):



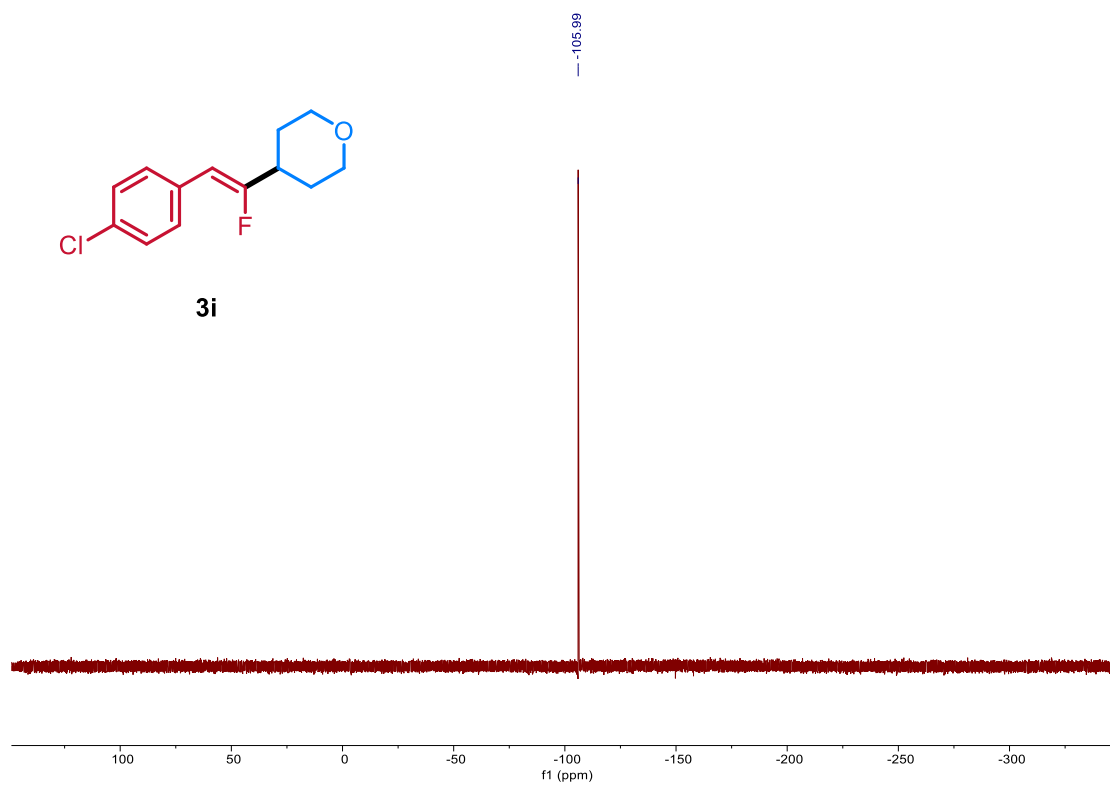
¹H NMR of Compound 3i (400 MHz, Chloroform-d):



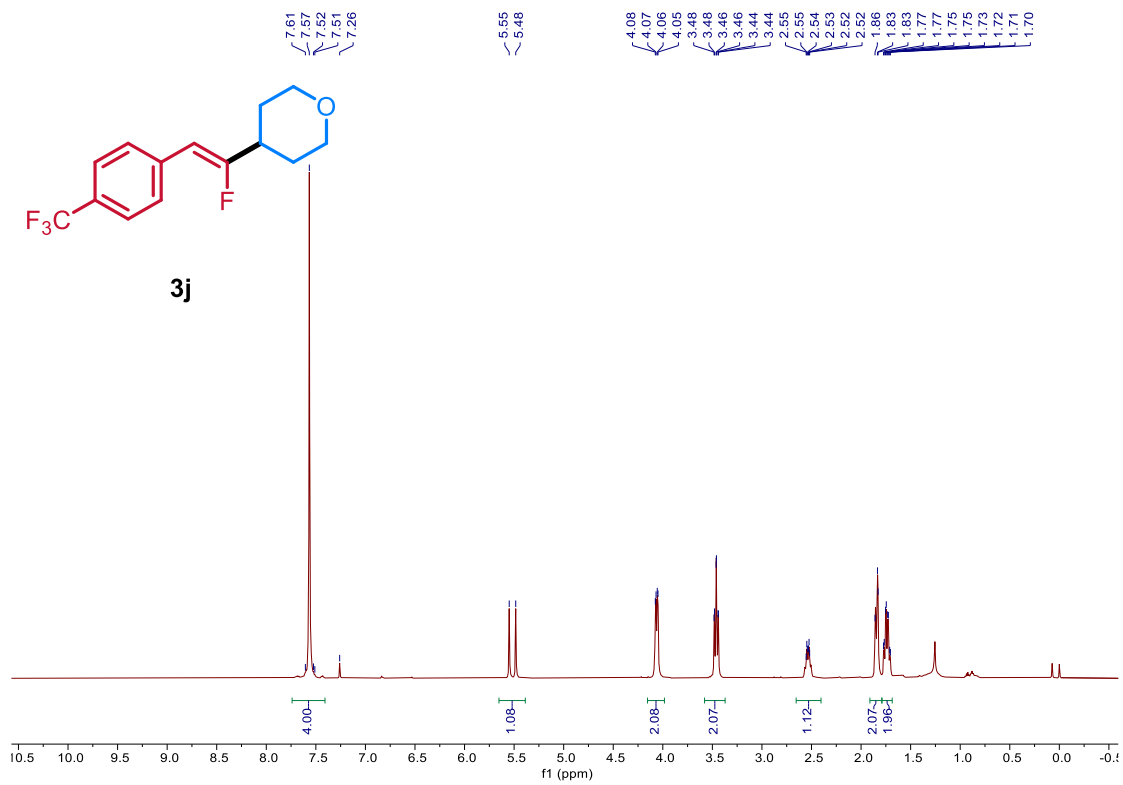
¹³C NMR of Compound 3i (150 MHz, Chloroform-d):



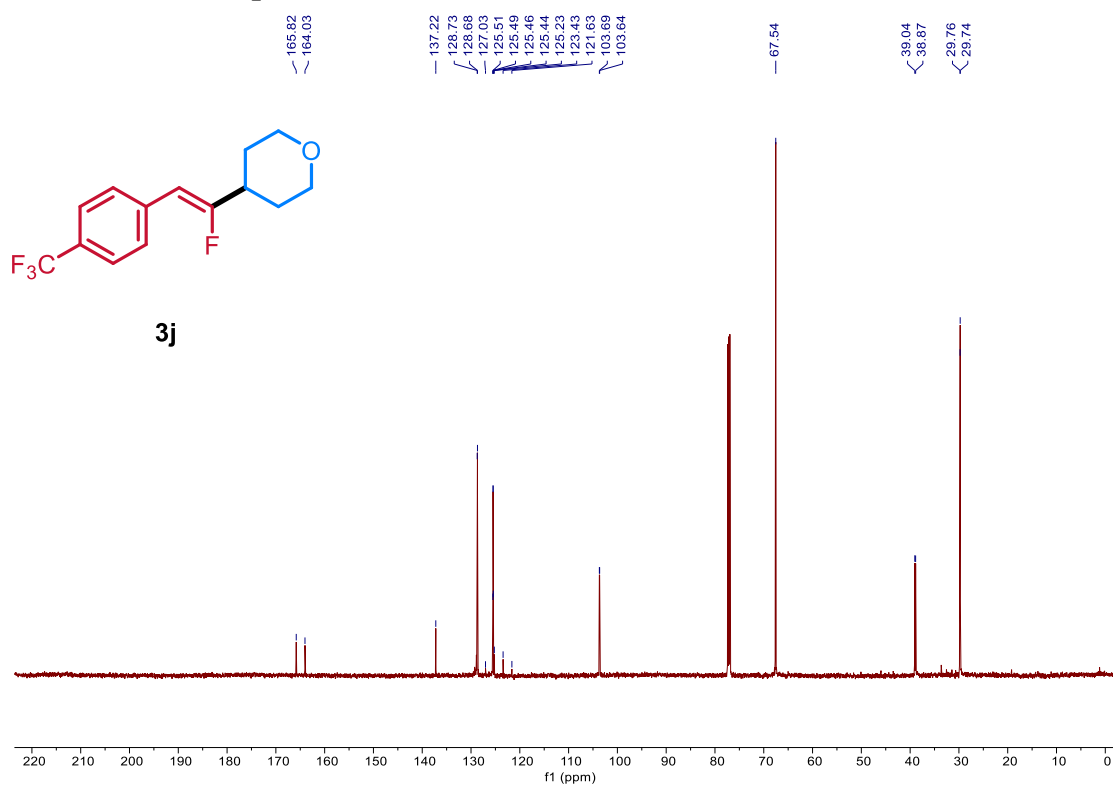
¹⁹F NMR of Compound 3i (375 MHz, Chloroform-*d*):



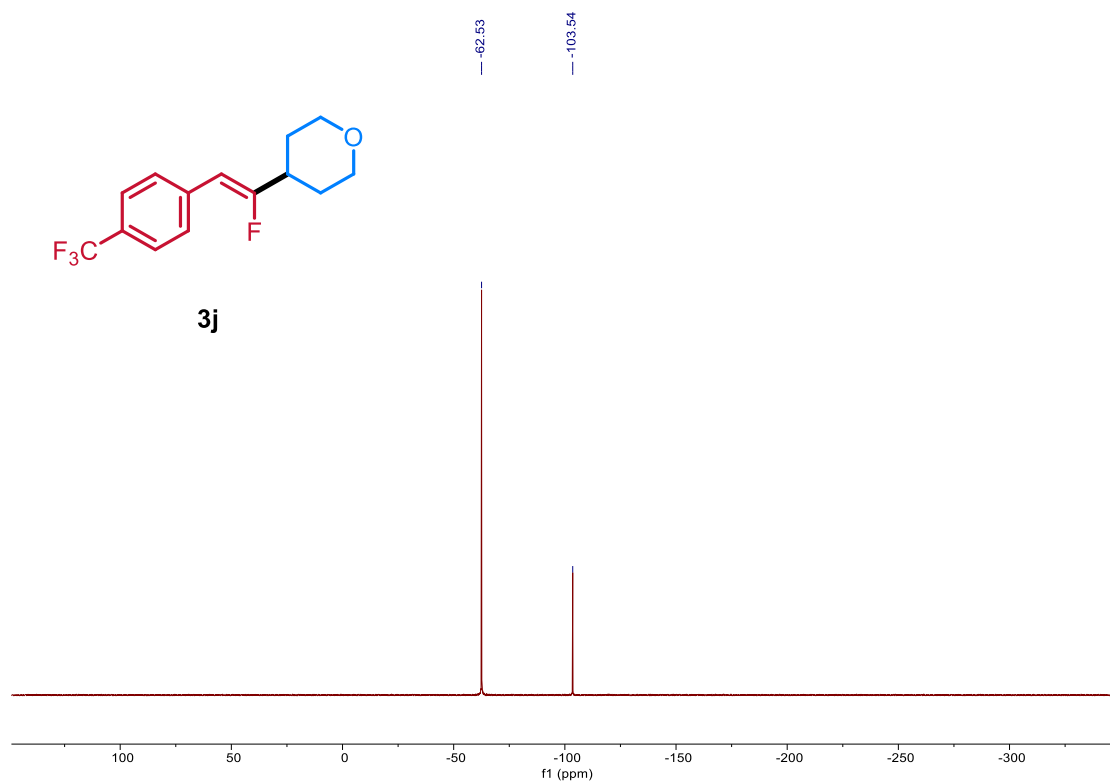
¹H NMR of Compound 3j (600 MHz, Chloroform-*d*):



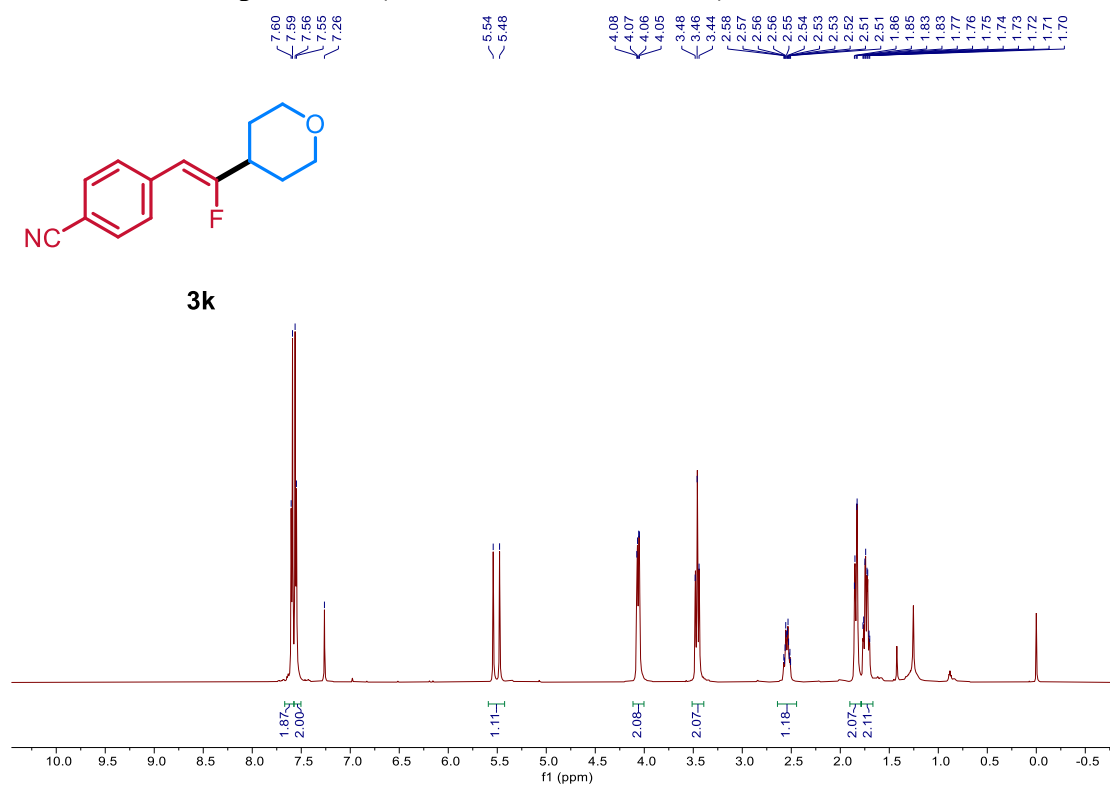
¹³C NMR of Compound 3i (150 MHz, Chloroform-*d*):



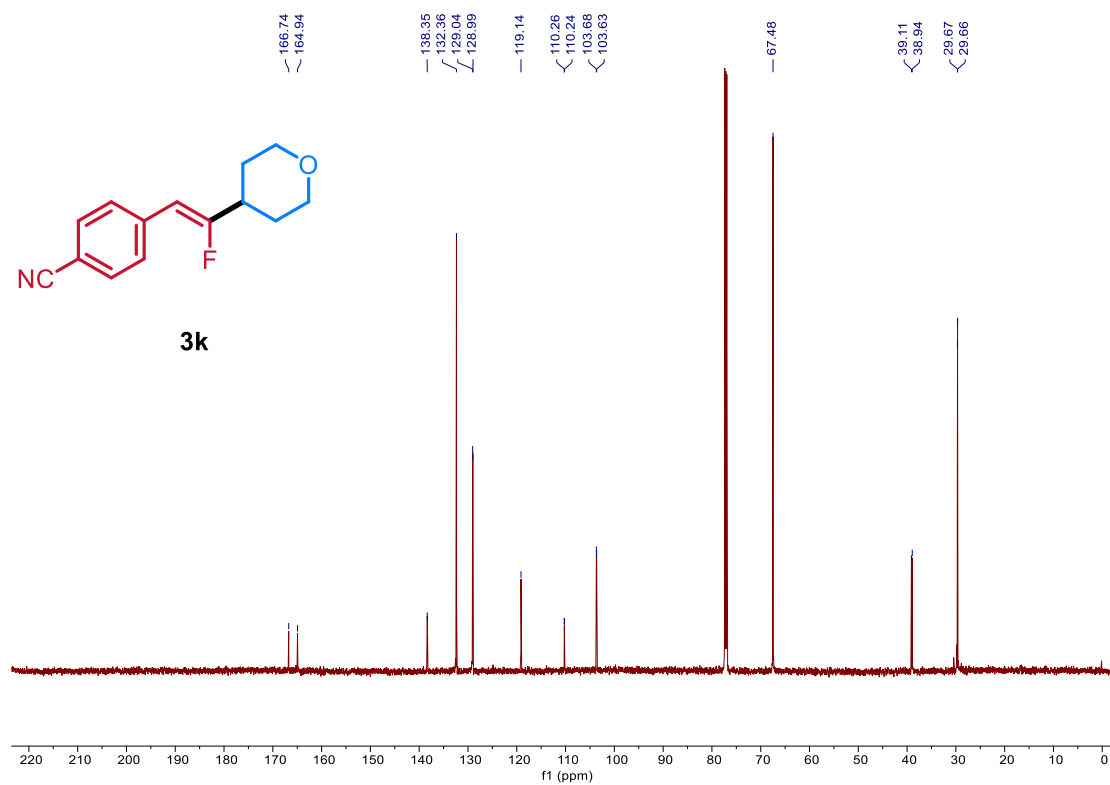
¹⁹F NMR of Compound 3j (375 MHz, Chloroform-*d*):



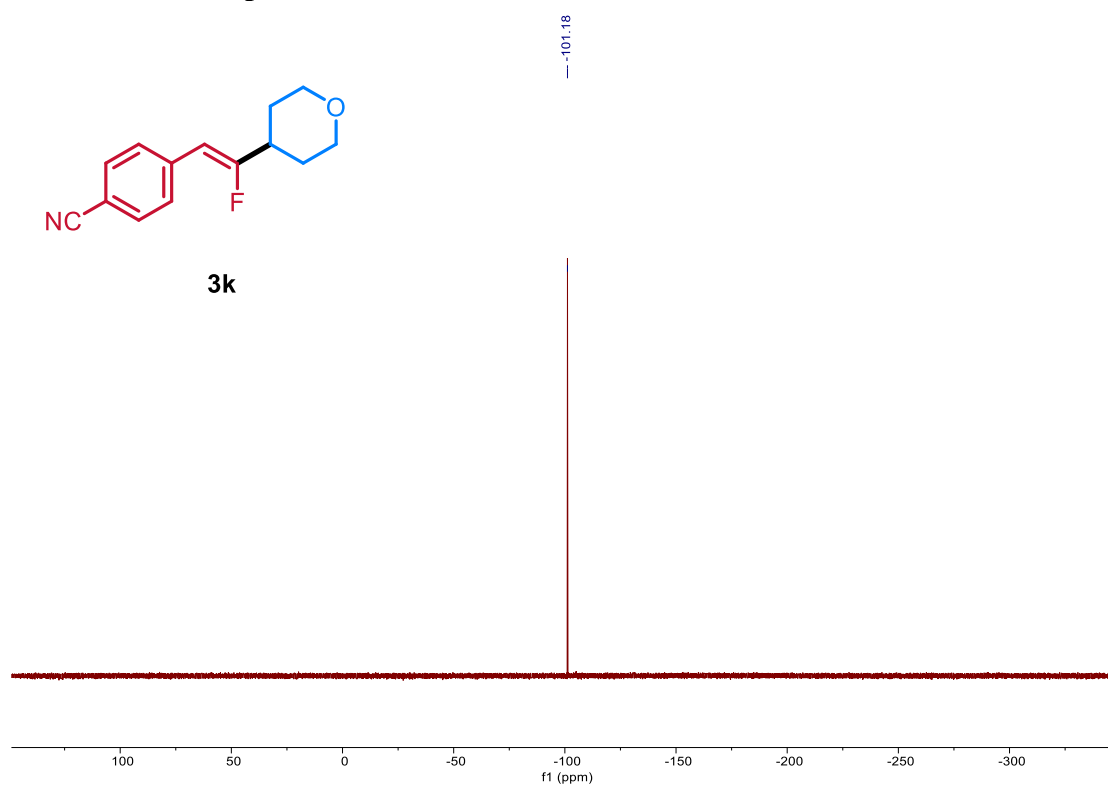
¹H NMR of Compound 3k (600 MHz, Chloroform-*d*):



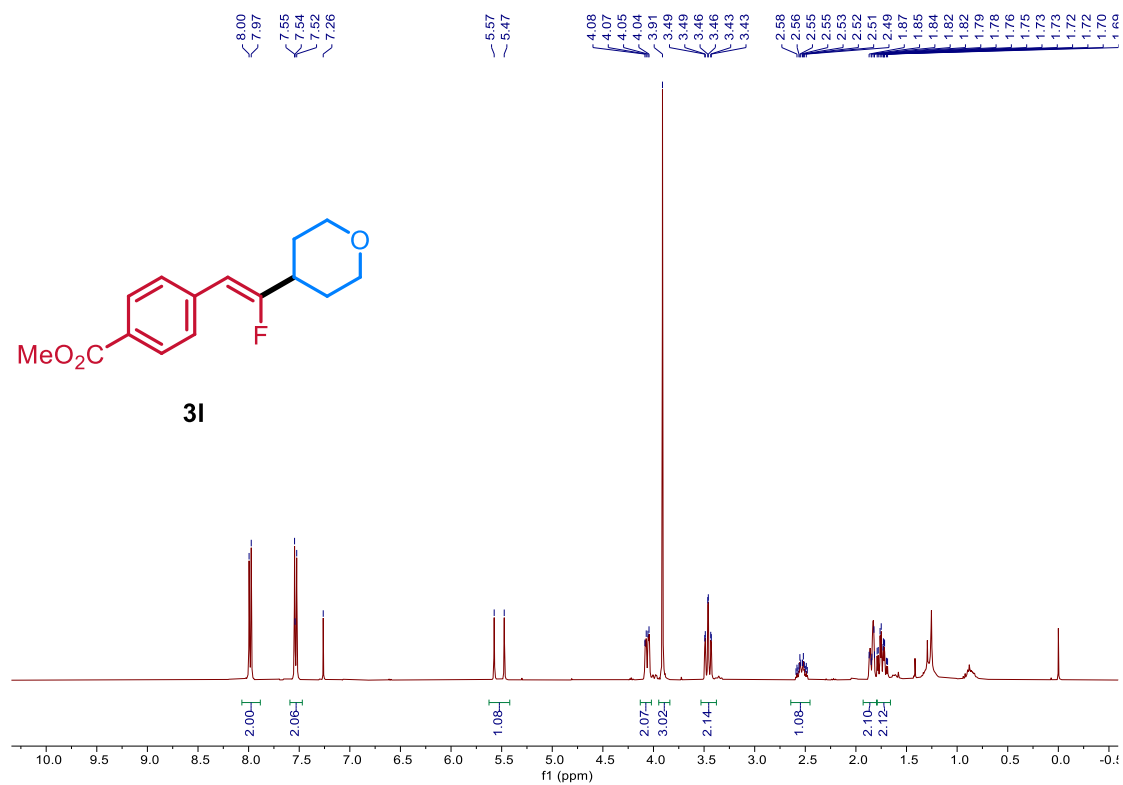
¹³C NMR of Compound 3k (150 MHz, Chloroform-*d*):



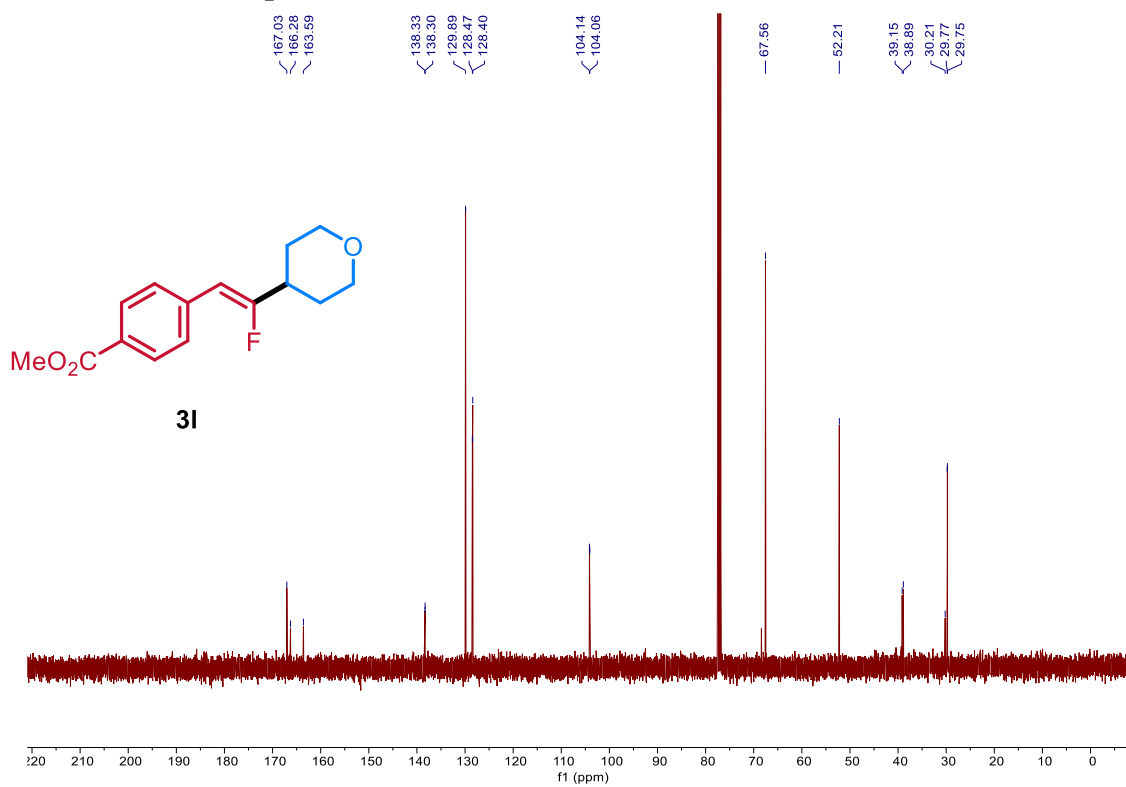
¹⁹F NMR of Compound 3k (375 MHz, Chloroform-*d*):



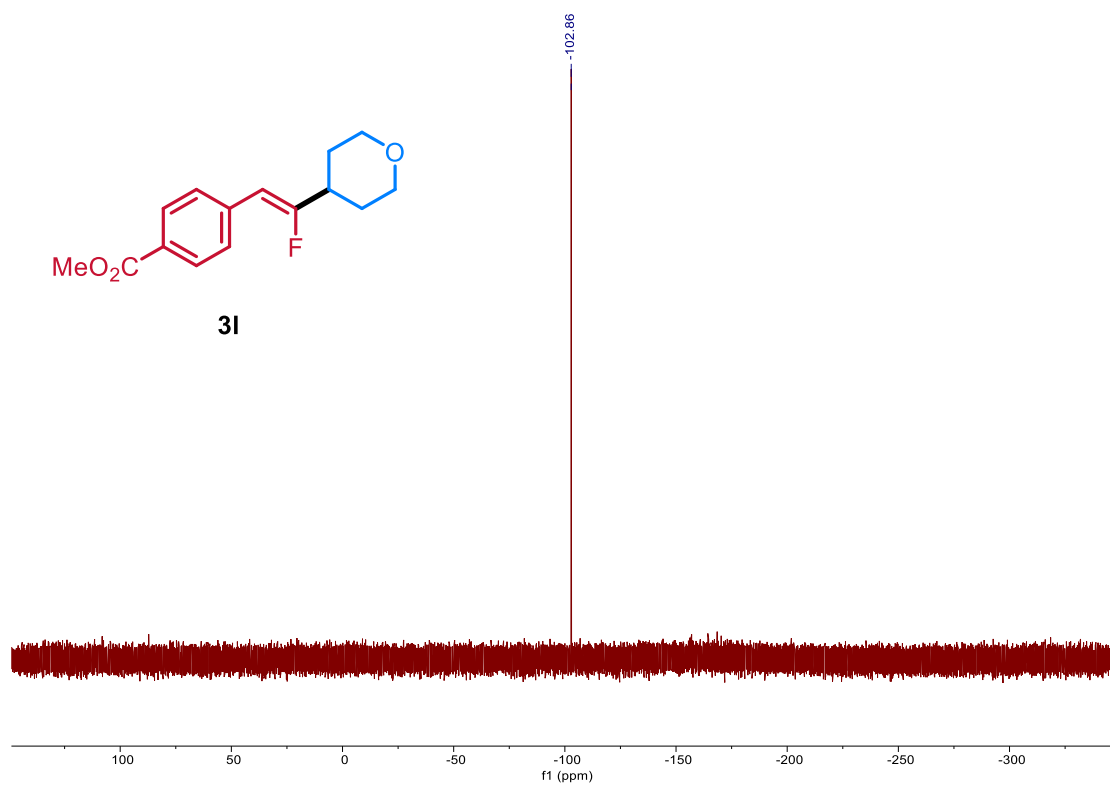
¹H NMR of Compound 3l (400 MHz, Chloroform-*d*):



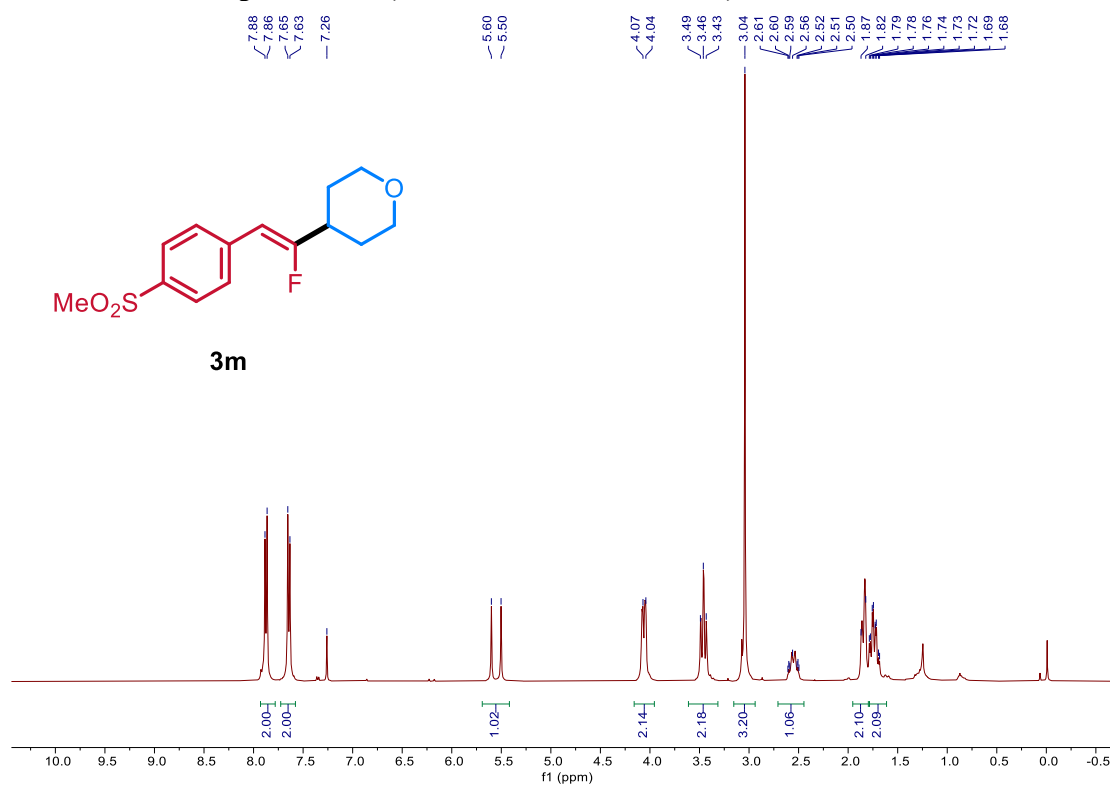
¹³C NMR of Compound 3I (100 MHz, Chloroform-*d*):



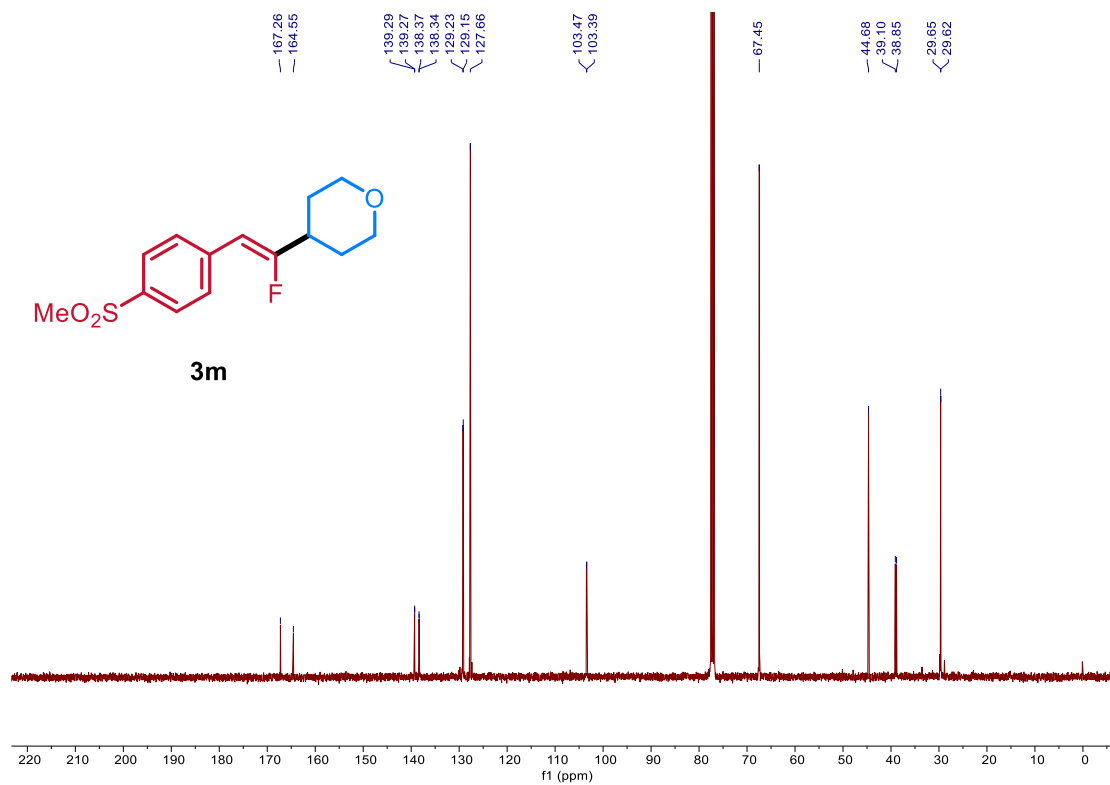
¹⁹F NMR of Compound 3I (375 MHz, Chloroform-*d*):



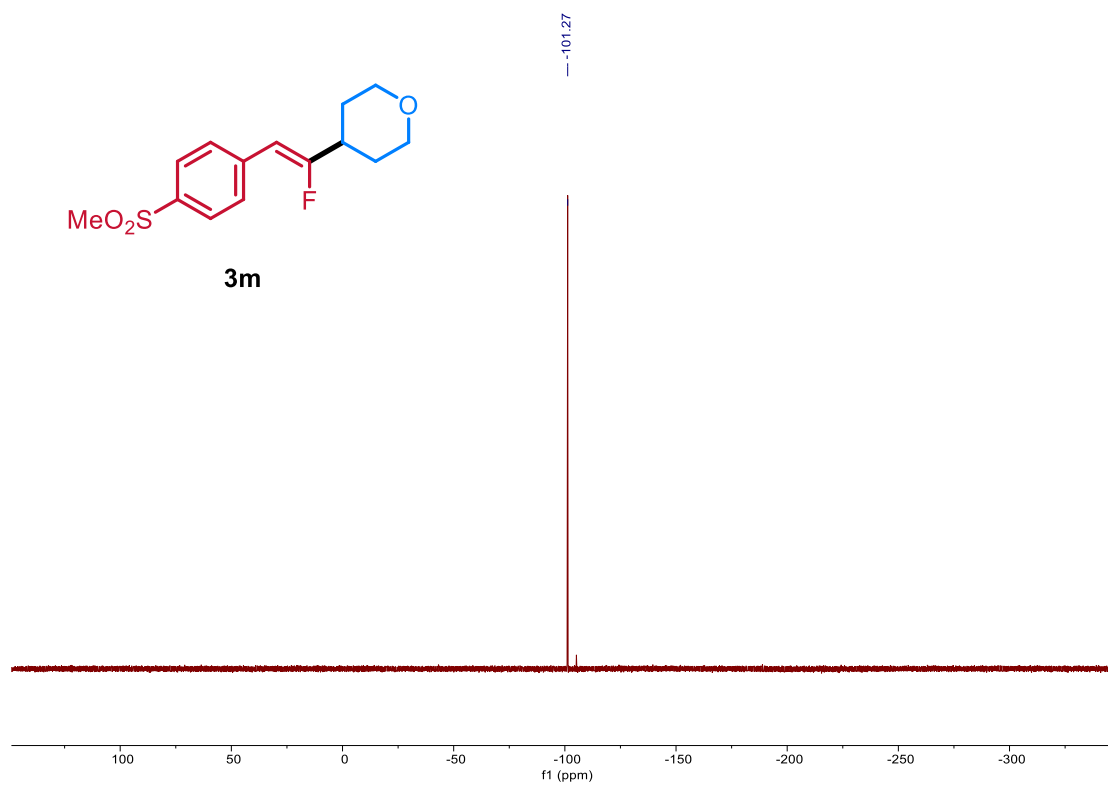
¹H NMR of Compound 3m (400 MHz, Chloroform-*d*):



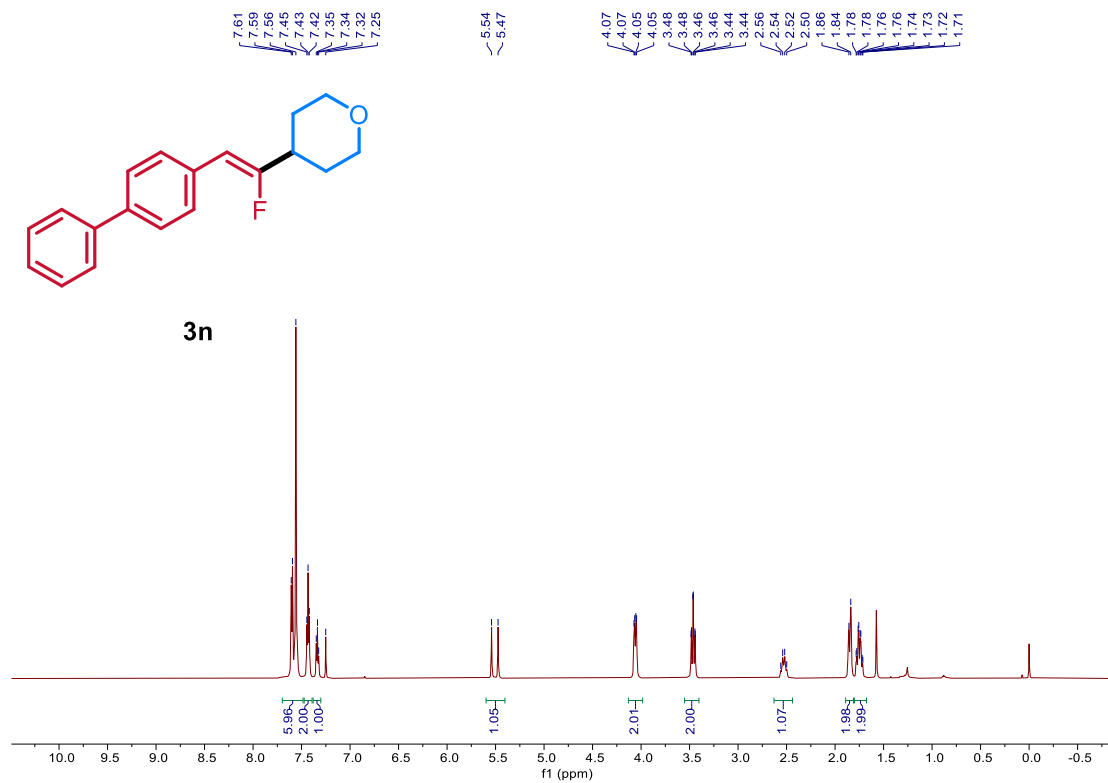
¹³C NMR of Compound 3m (100 MHz, Chloroform-*d*):



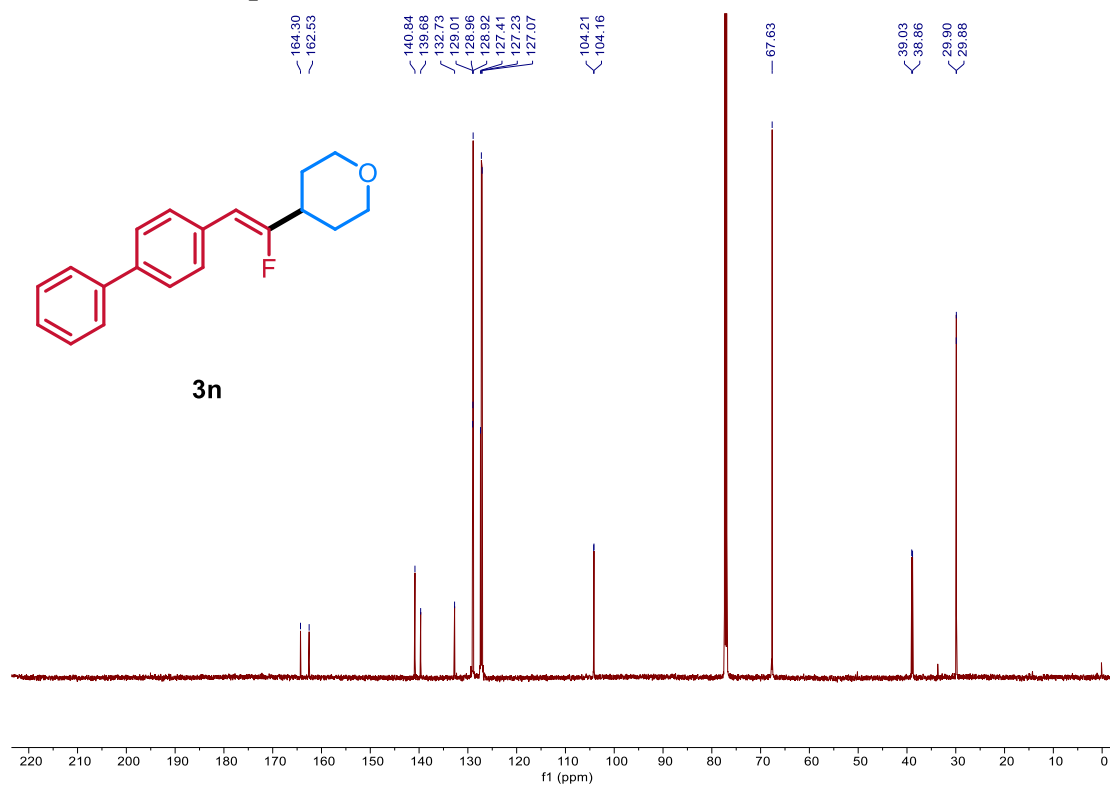
^{19}F NMR of Compound 3m (375 MHz, Chloroform-*d*):



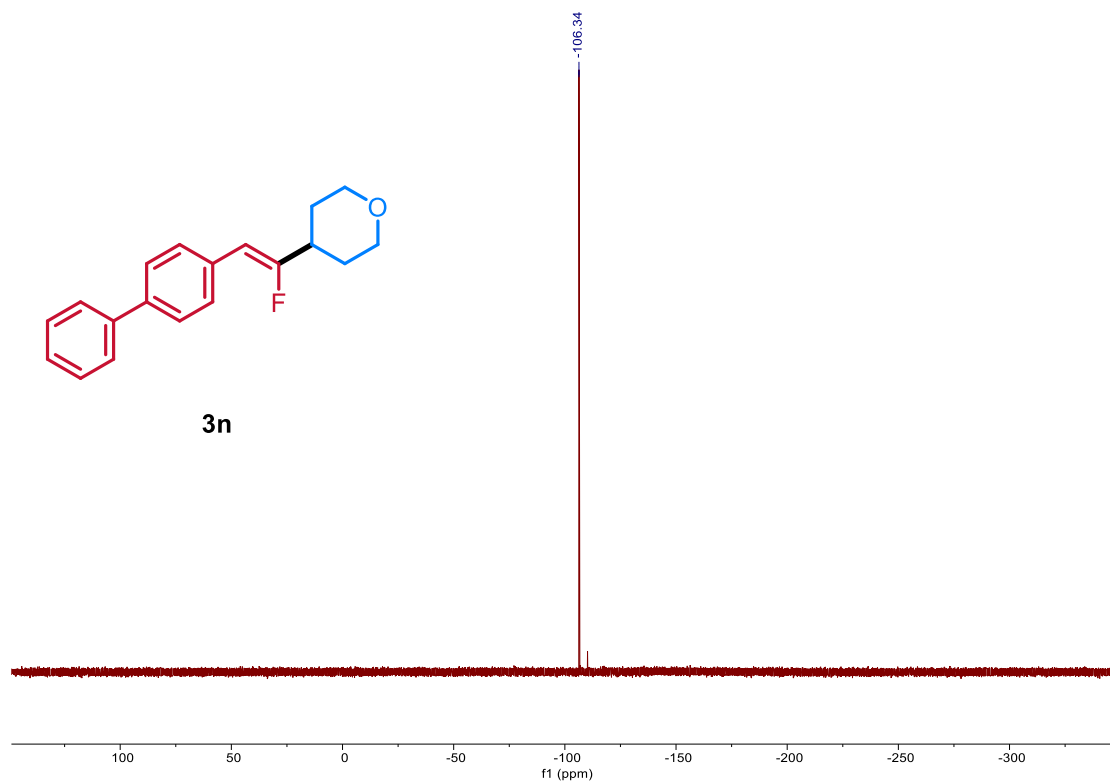
^1H NMR of Compound 3n (600 MHz, Chloroform-*d*):



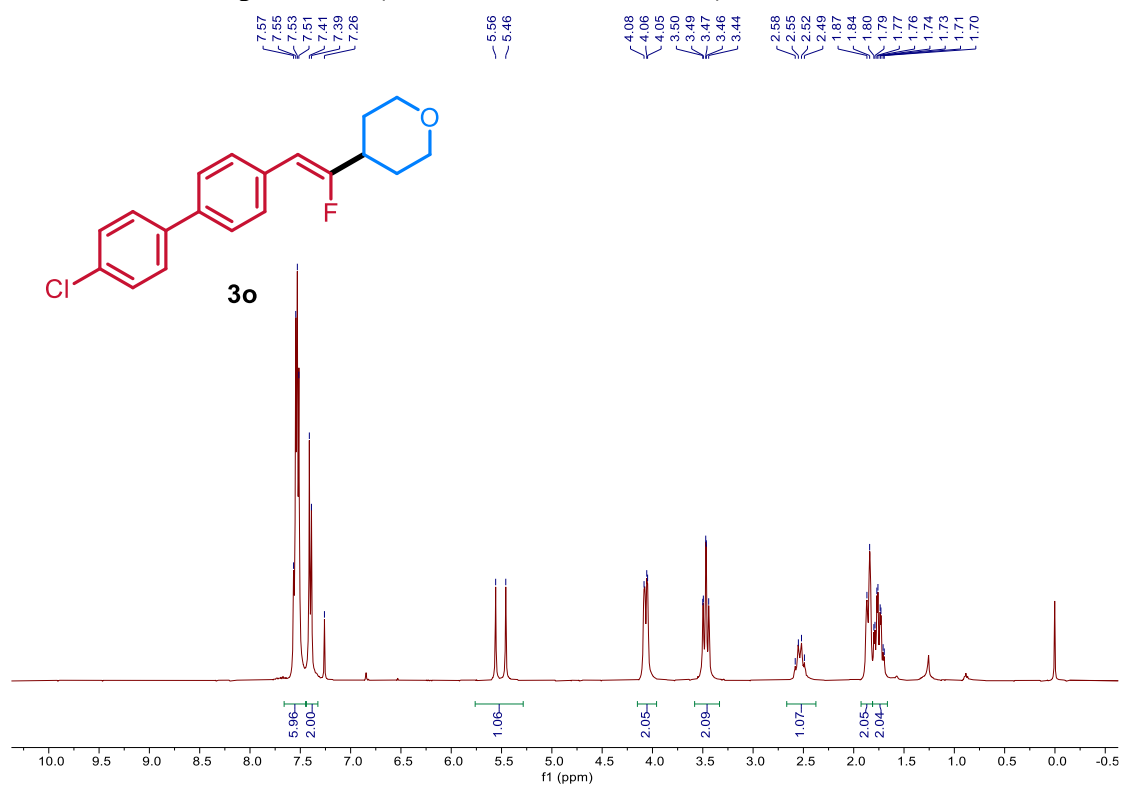
¹³C NMR of Compound 3n (150 MHz, Chloroform-*d*):



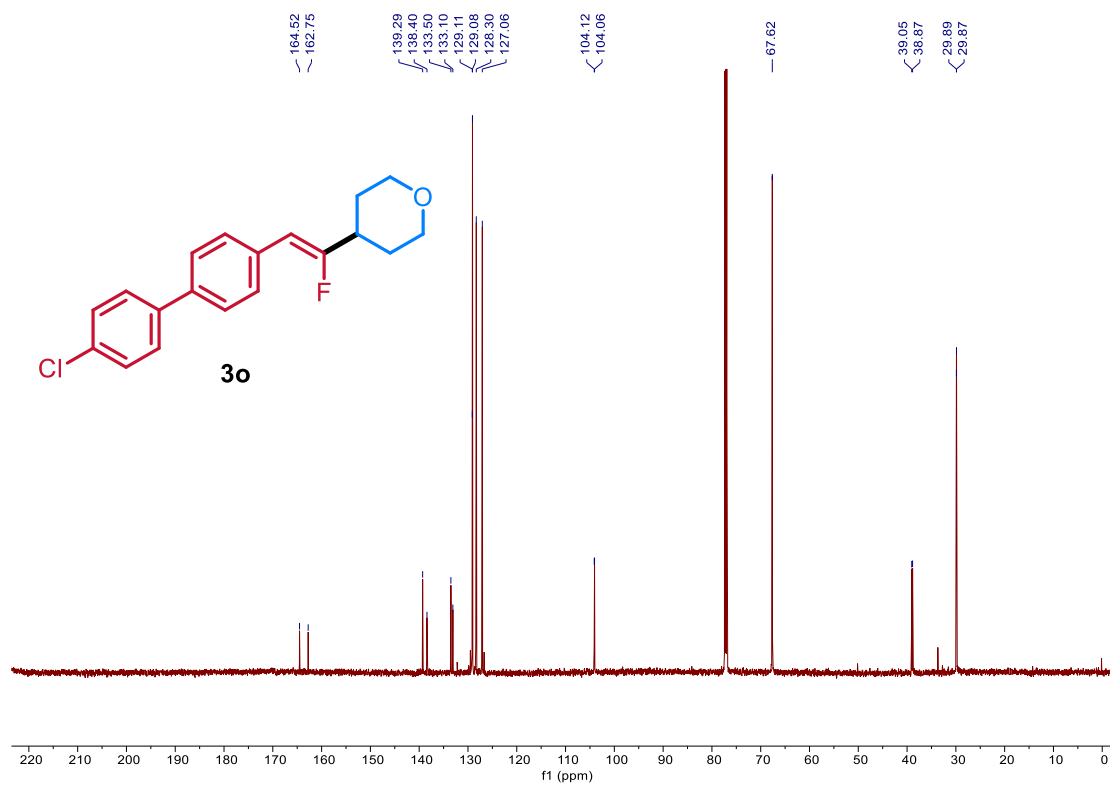
¹⁹F NMR of Compound 3n (375 MHz, Chloroform-*d*):



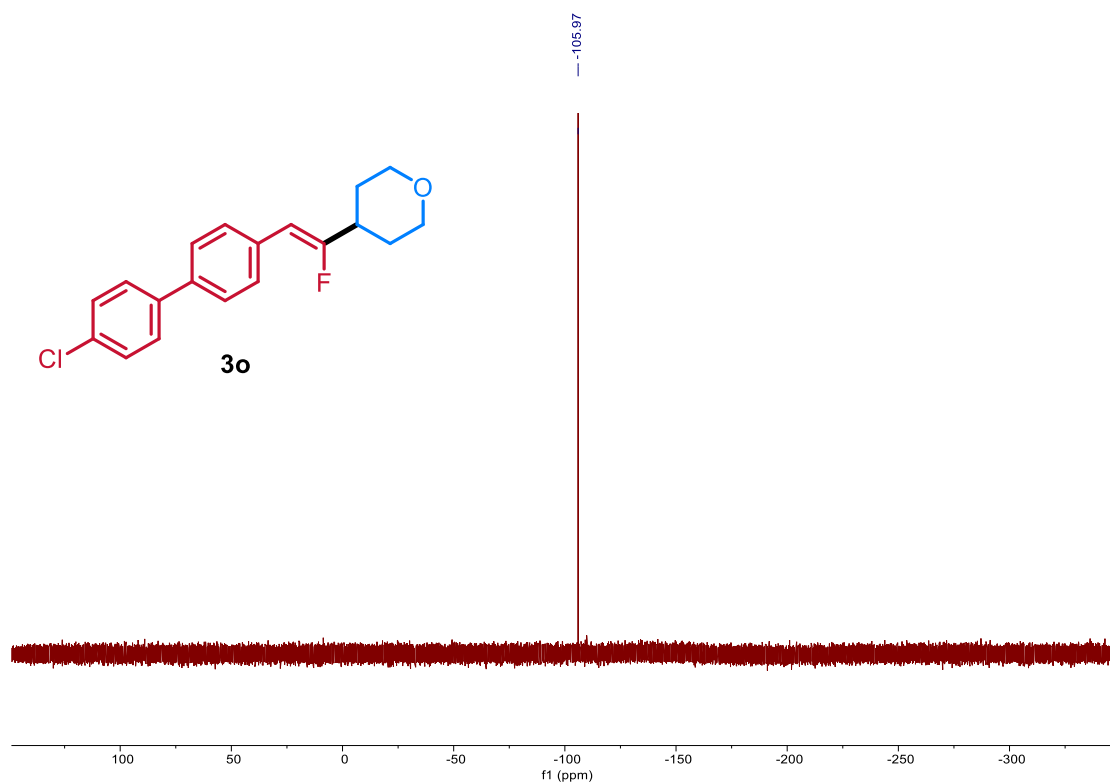
¹H NMR of Compound 3o (600 MHz, Chloroform-*d*):



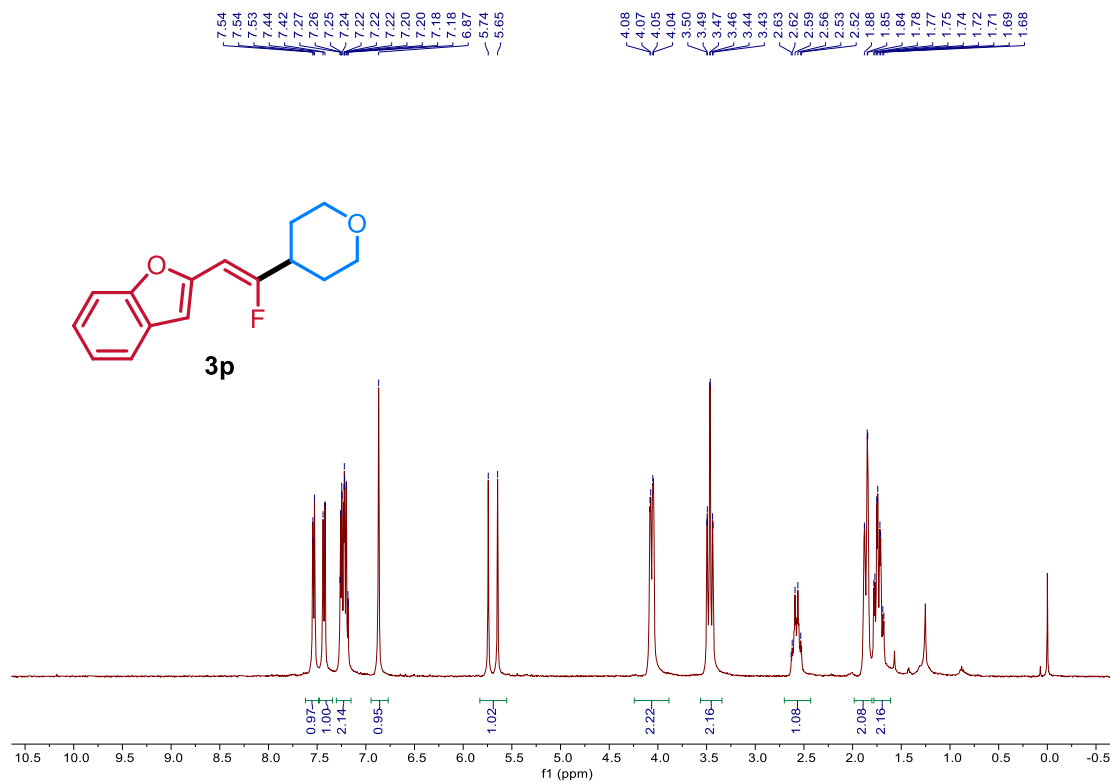
¹³C NMR of Compound 3o (150 MHz, Chloroform-*d*):



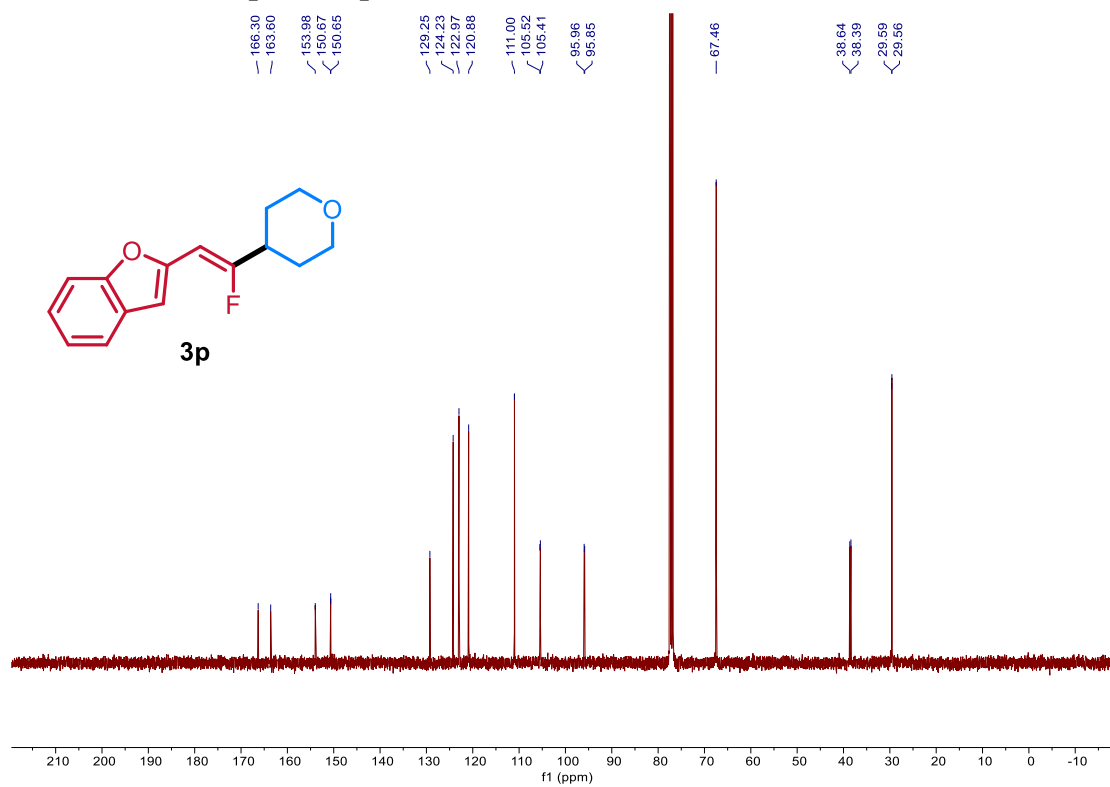
¹⁹F NMR of Compound 3o (375 MHz, Chloroform-*d*):



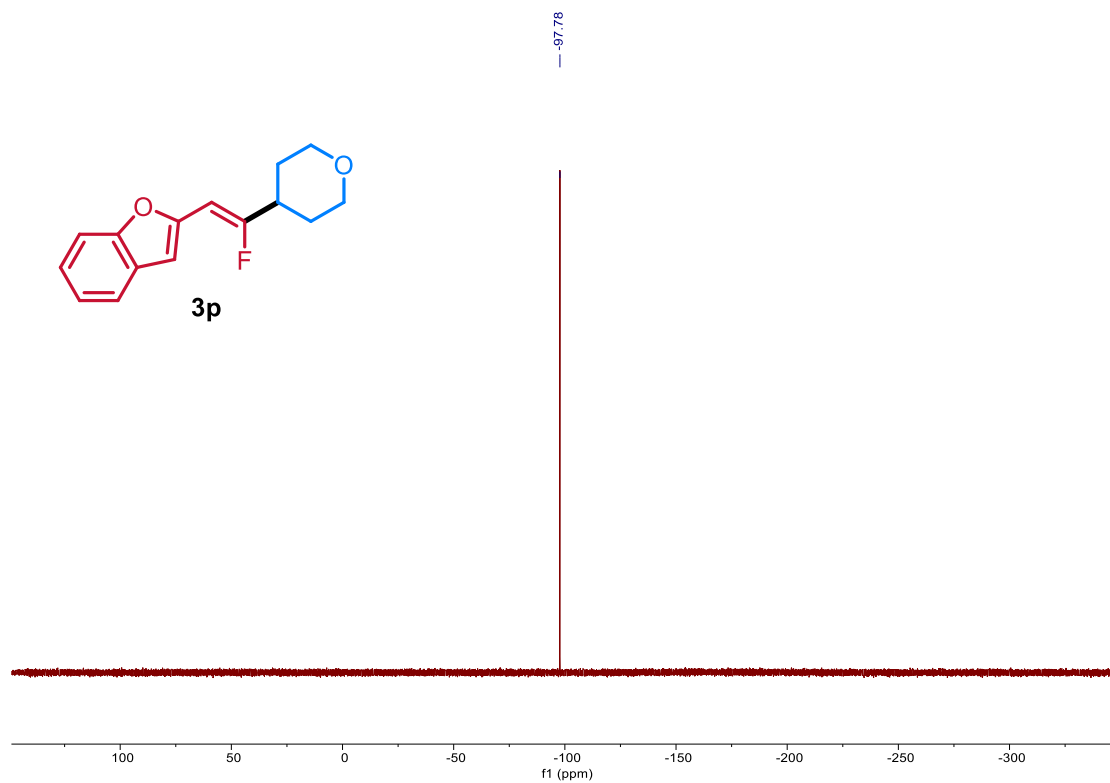
¹H NMR of Compound 3p (400 MHz, Chloroform-*d*):



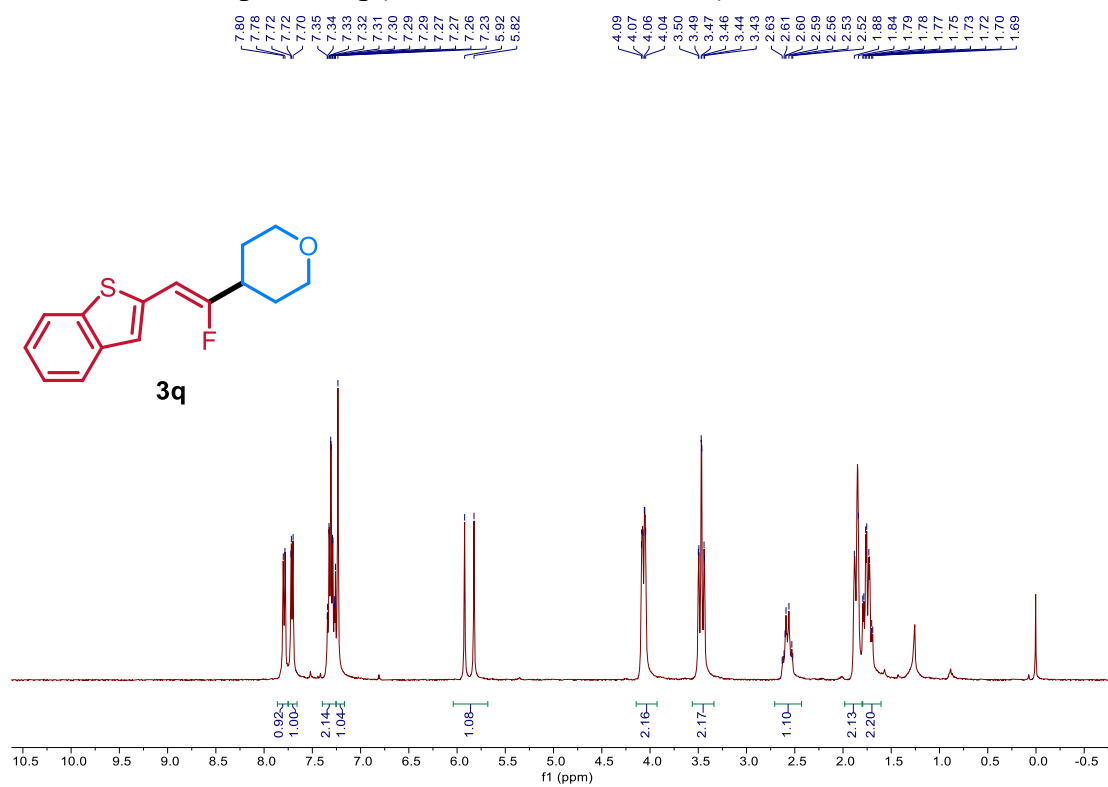
¹³C NMR of Compound 3p (100 MHz, Chloroform-*d*):



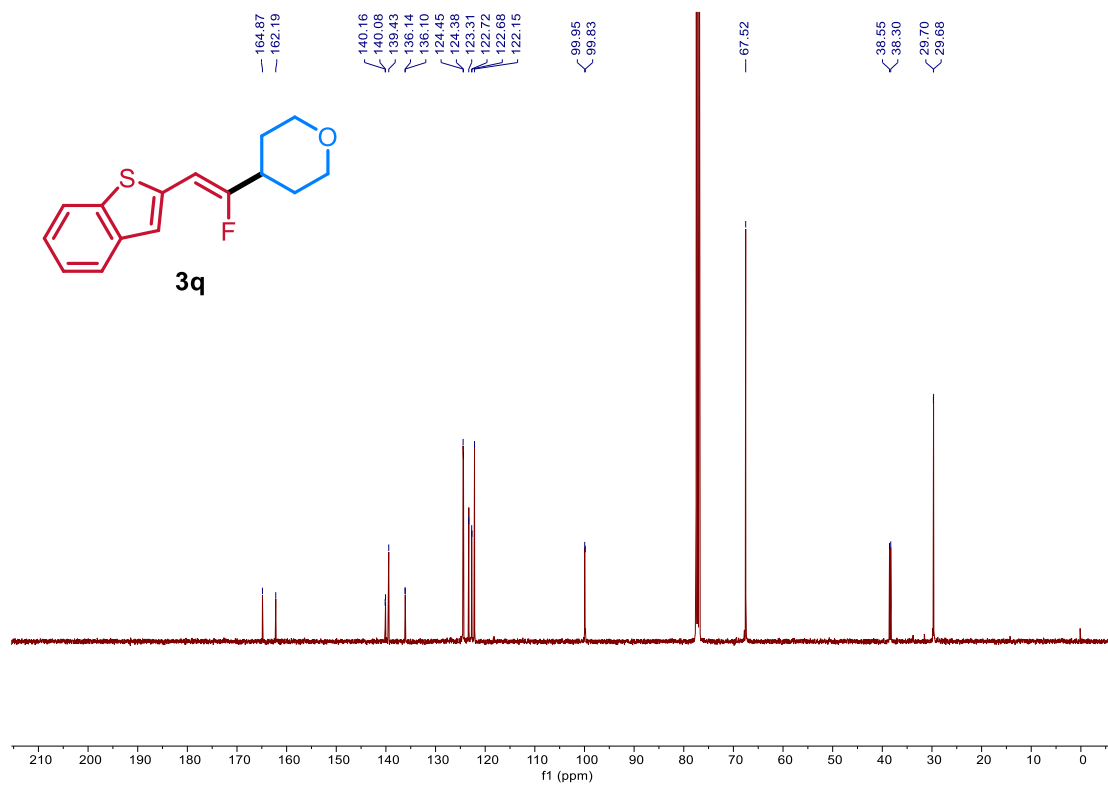
¹⁹F NMR of Compound 3p (375 MHz, Chloroform-*d*):



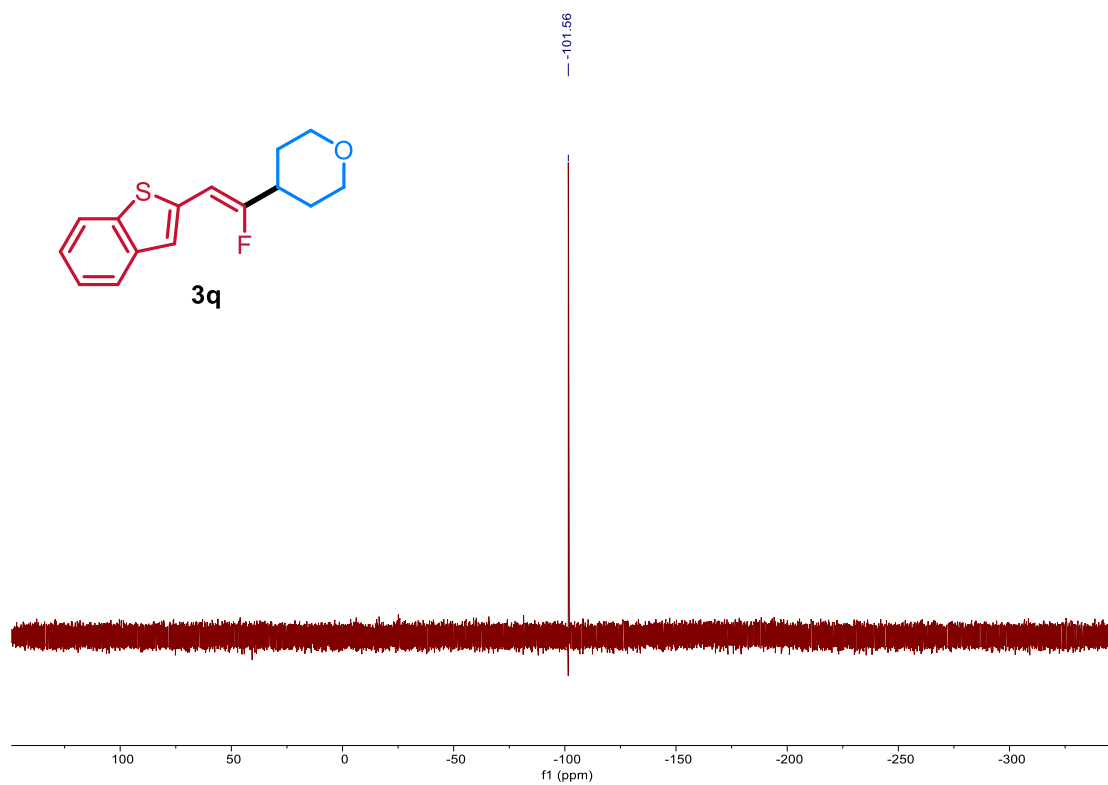
¹H NMR of Compound 3q (400 MHz, Chloroform-*d*):



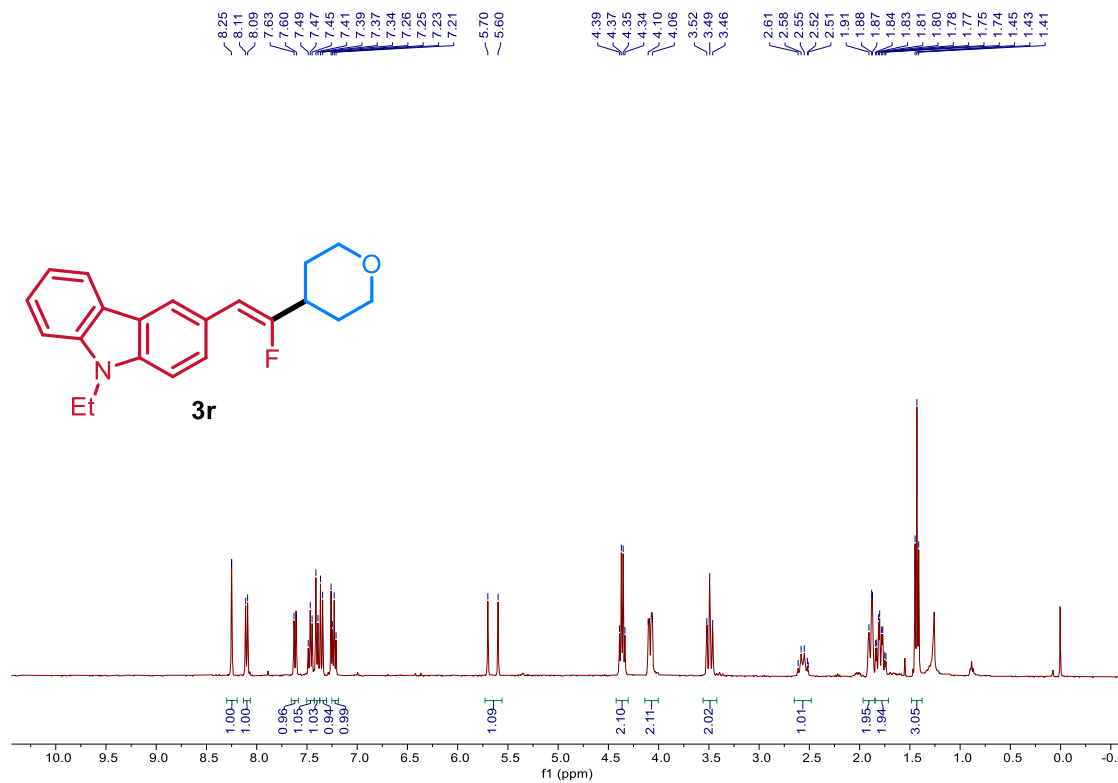
¹³C NMR of Compound 3q (100 MHz, Chloroform-*d*):



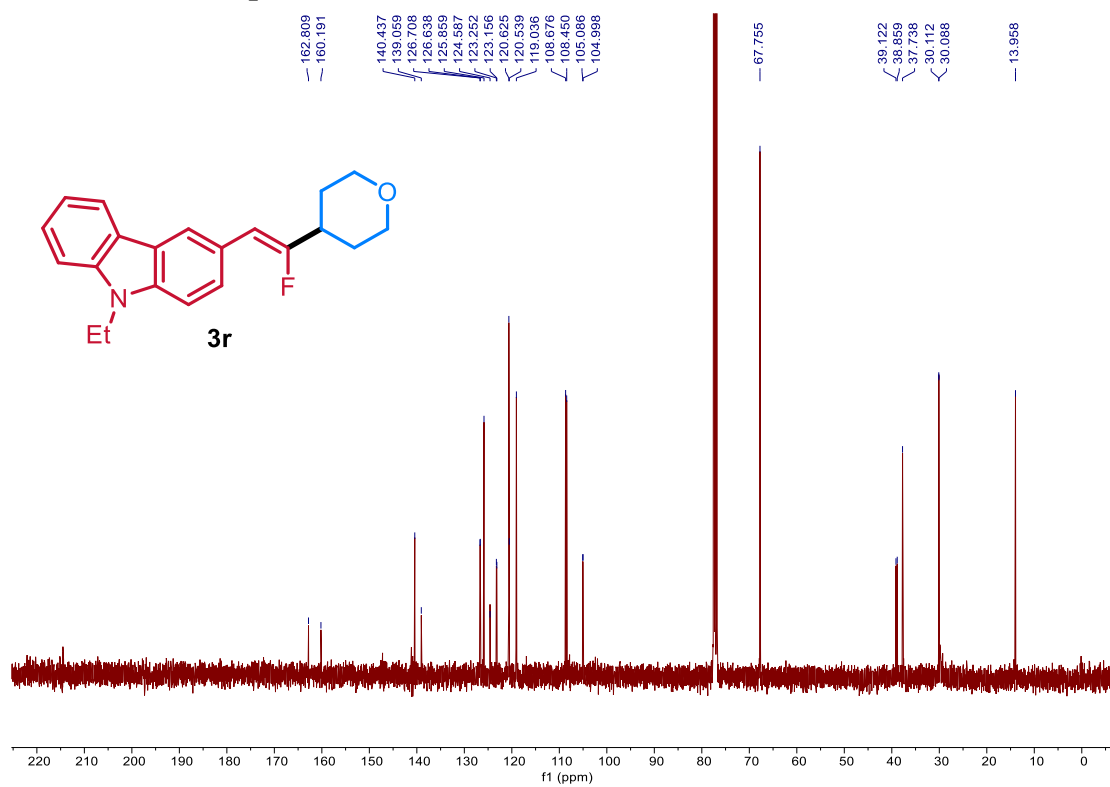
¹⁹F NMR of Compound 3q (375 MHz, Chloroform-*d*):



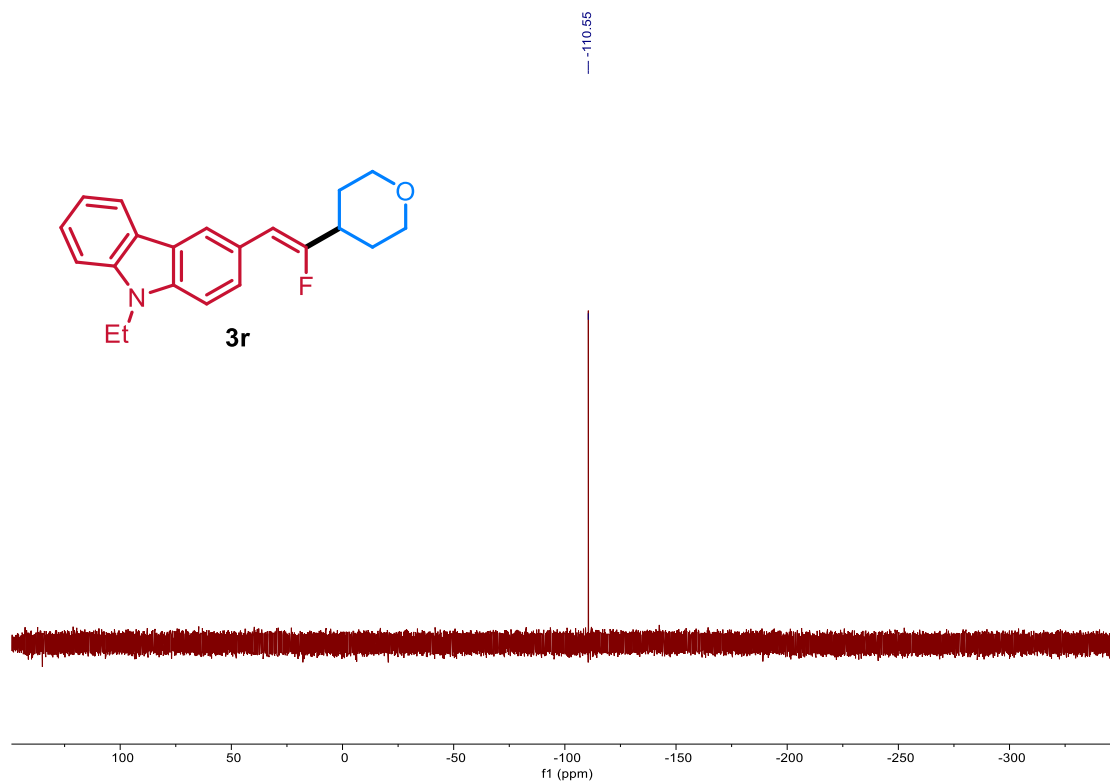
¹H NMR of Compound 3r (400 MHz, Chloroform-*d*):



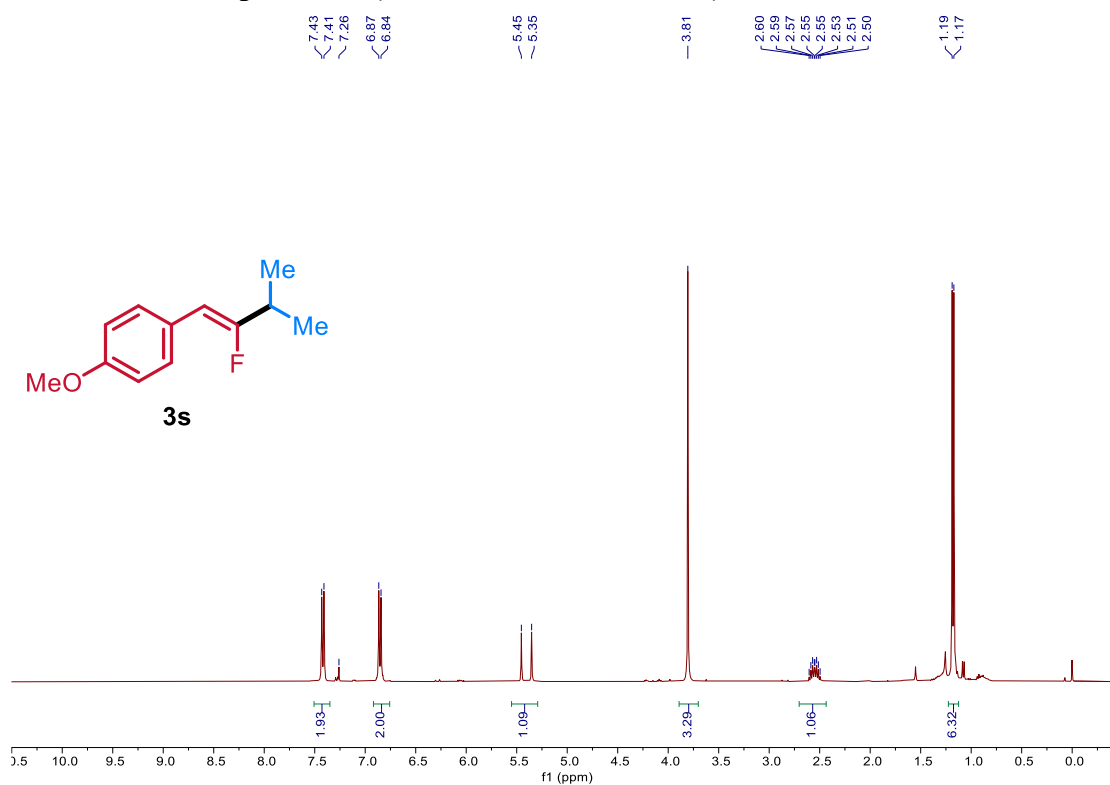
¹³C NMR of Compound 3r (100 MHz, Chloroform-*d*):



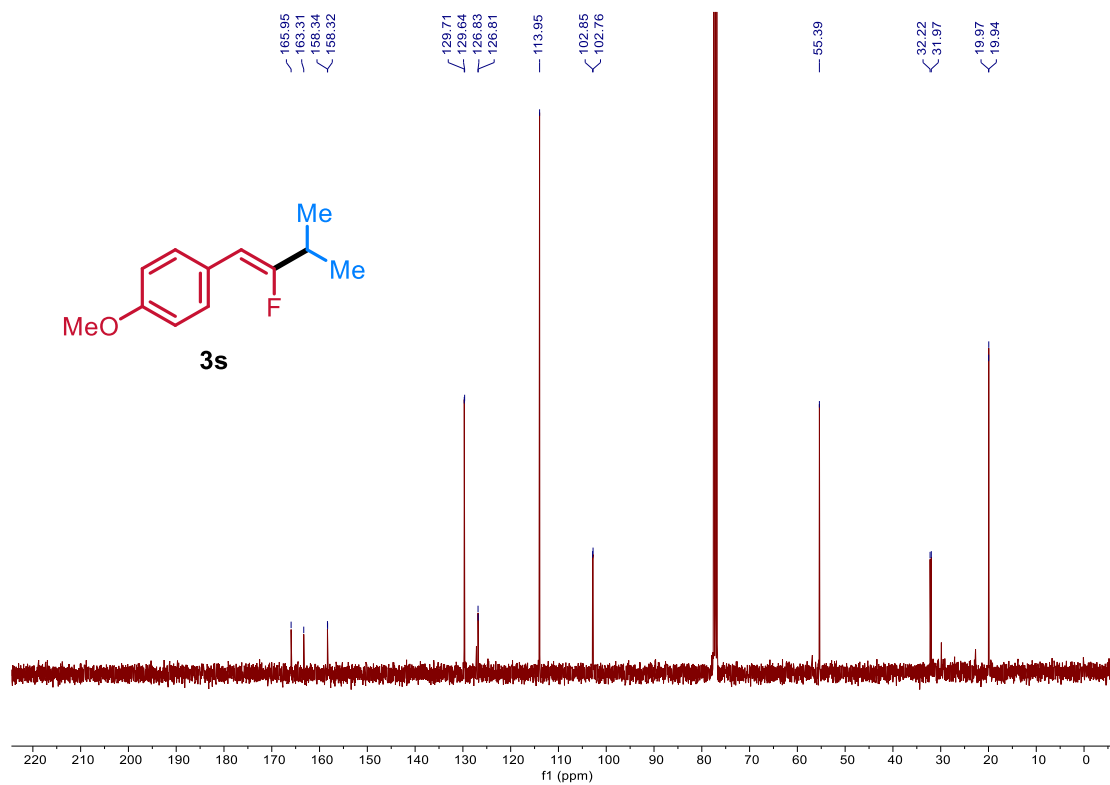
¹⁹F NMR of Compound 3r (375 MHz, Chloroform-*d*):



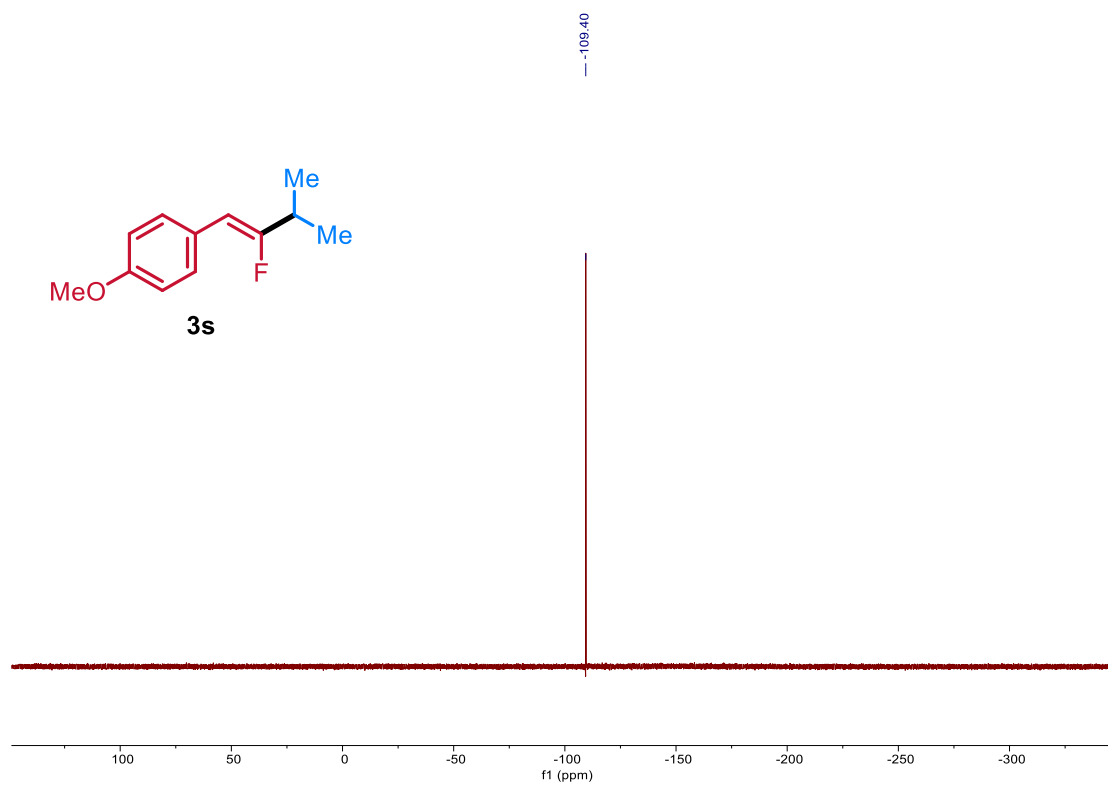
¹H NMR of Compound 3s (400 MHz, Chloroform-*d*):



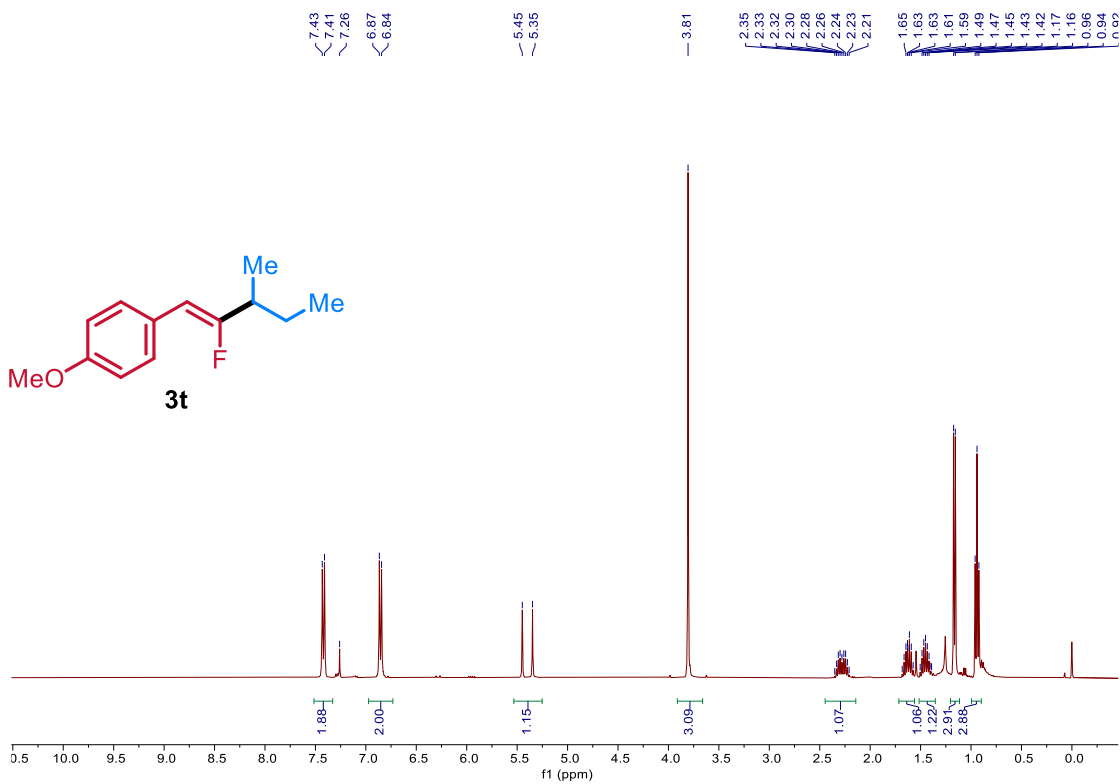
¹³C NMR of Compound 3s (100 MHz, Chloroform-*d*):



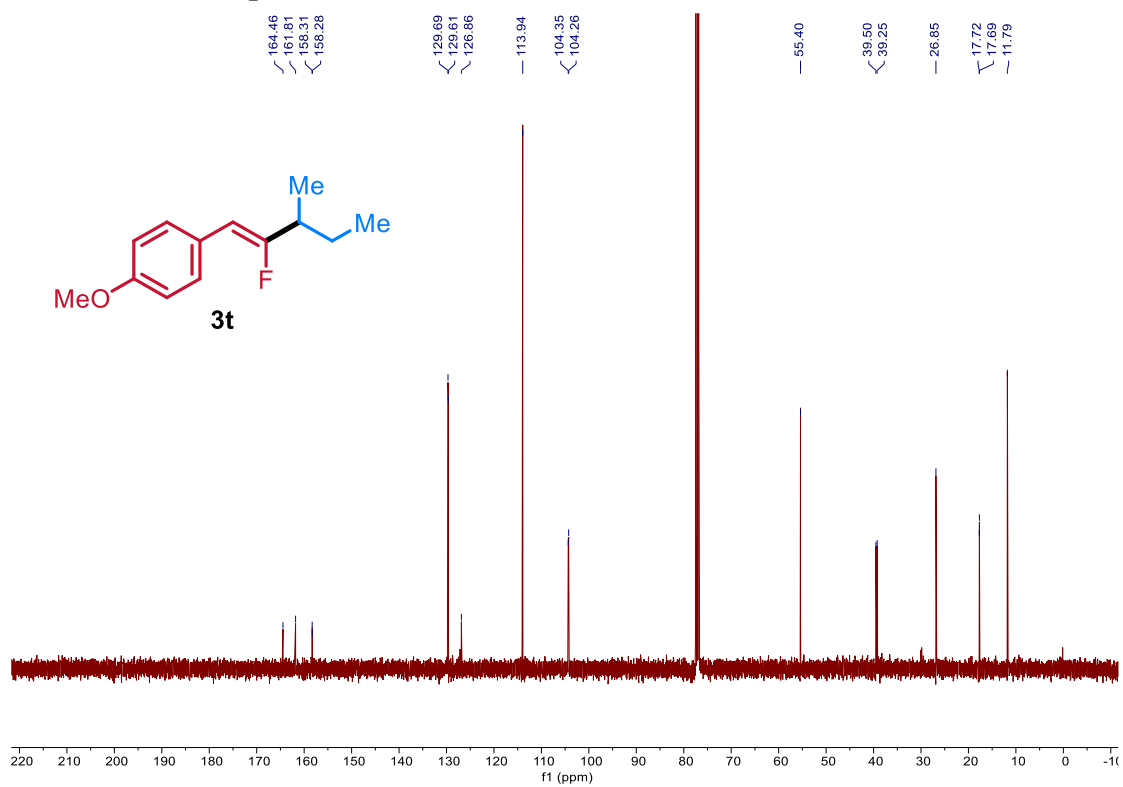
¹⁹F NMR of Compound 3s (375 MHz, Chloroform-*d*):



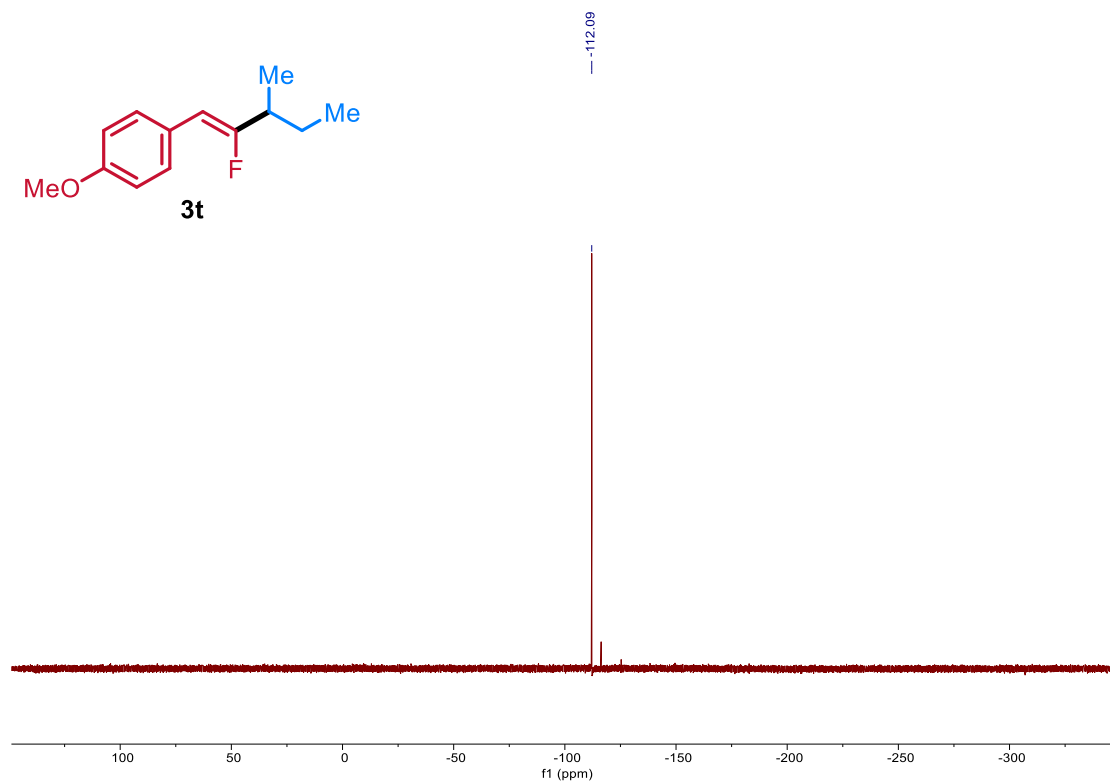
¹H NMR of Compound 3t (400 MHz, Chloroform-*d*):



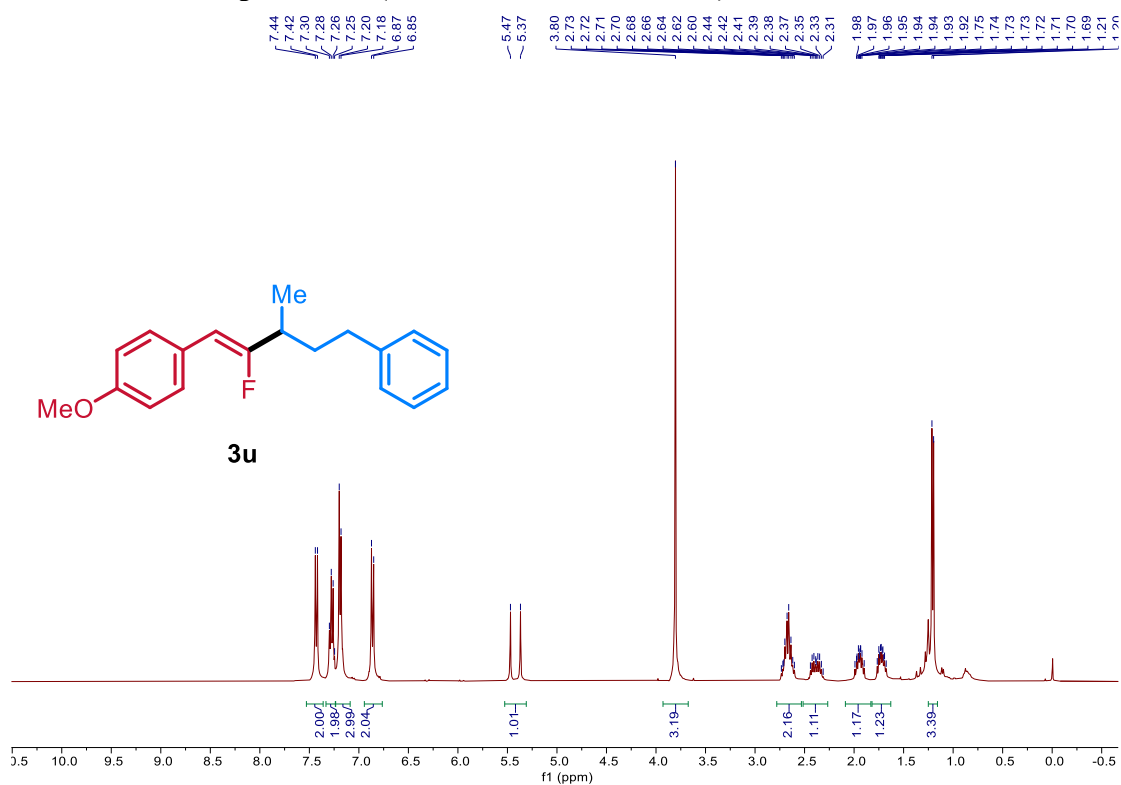
¹³C NMR of Compound 3t (100 MHz, Chloroform-*d*):



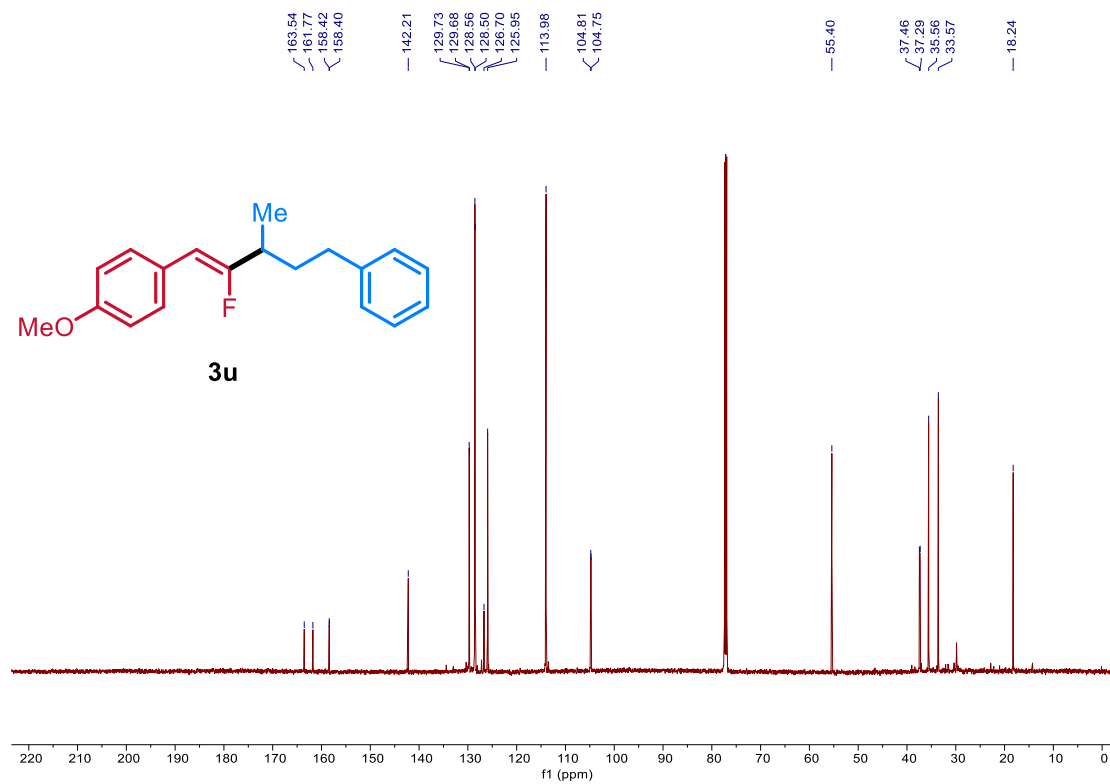
¹⁹F NMR of Compound 3t (375 MHz, Chloroform-*d*):



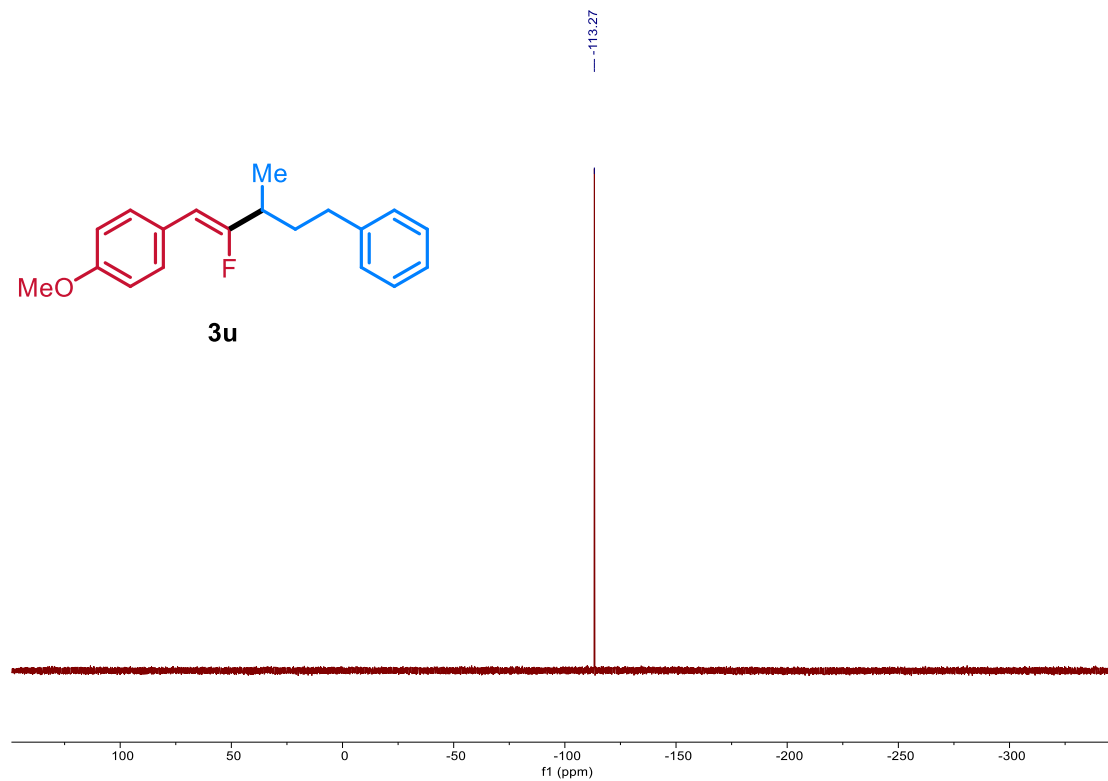
¹H NMR of Compound 3u (400 MHz, Chloroform-*d*):



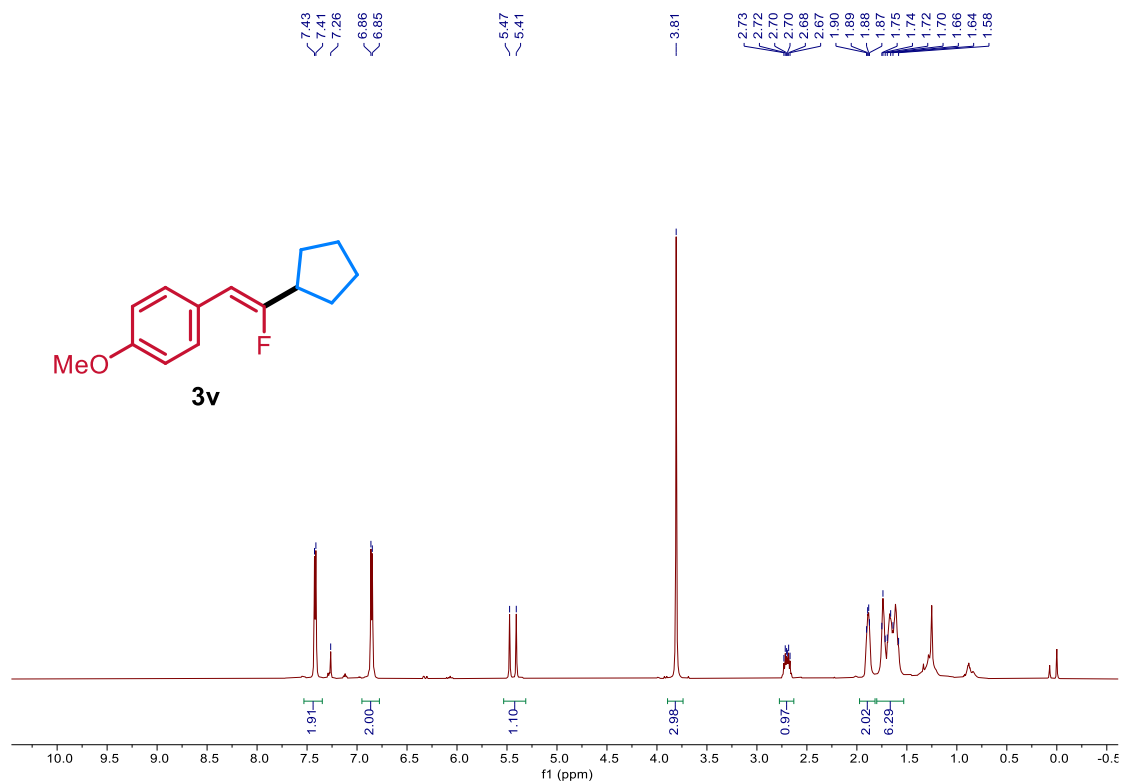
¹³C NMR of Compound 3u (150 MHz, Chloroform-*d*):



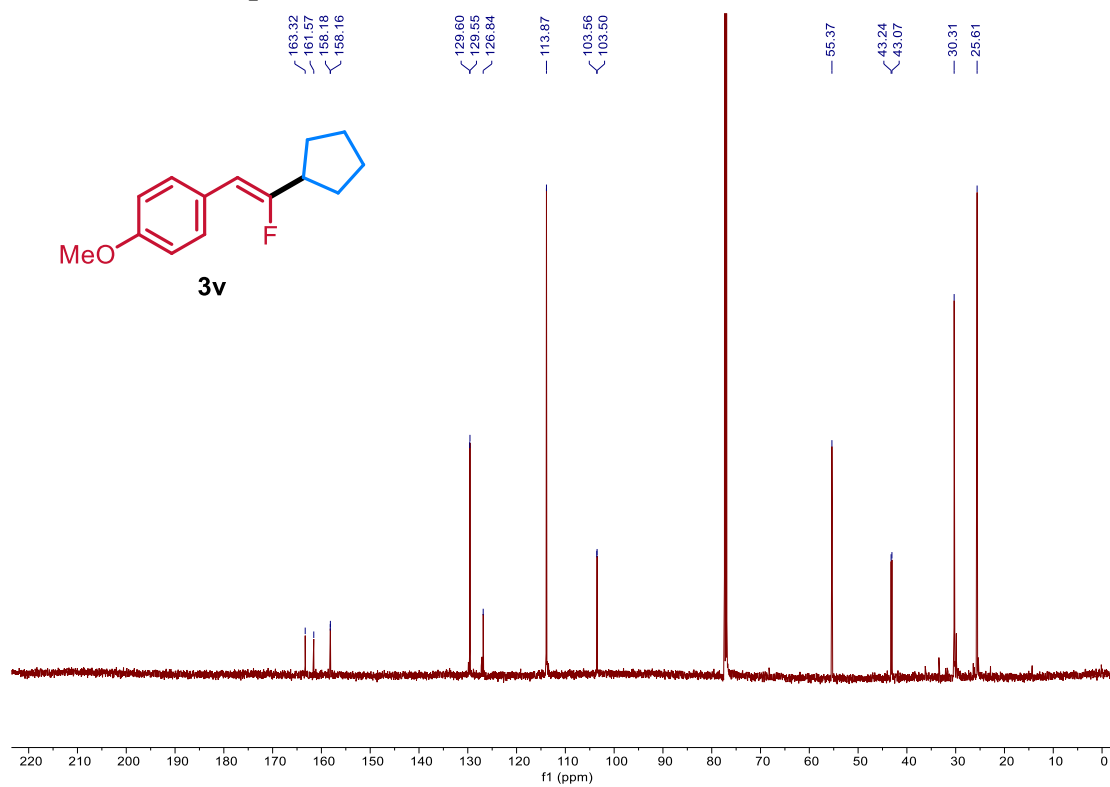
^{19}F NMR of Compound 3u (375 MHz, Chloroform-*d*):



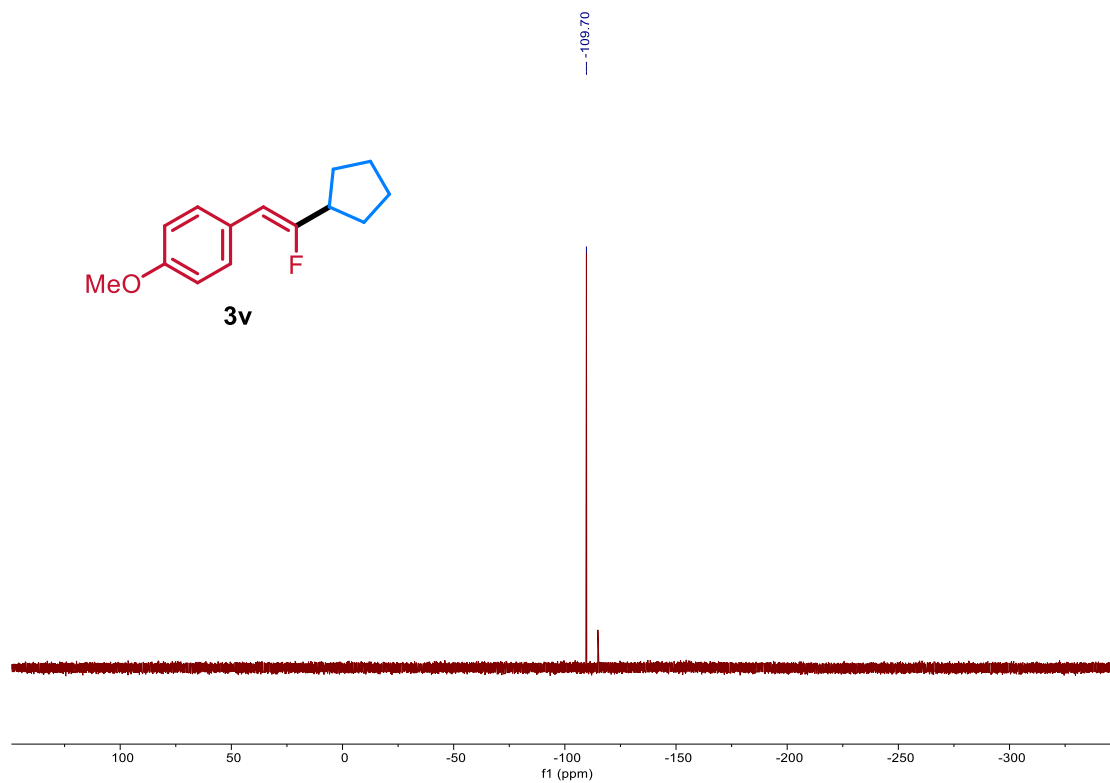
^1H NMR of Compound 3v (600 MHz, Chloroform-*d*):



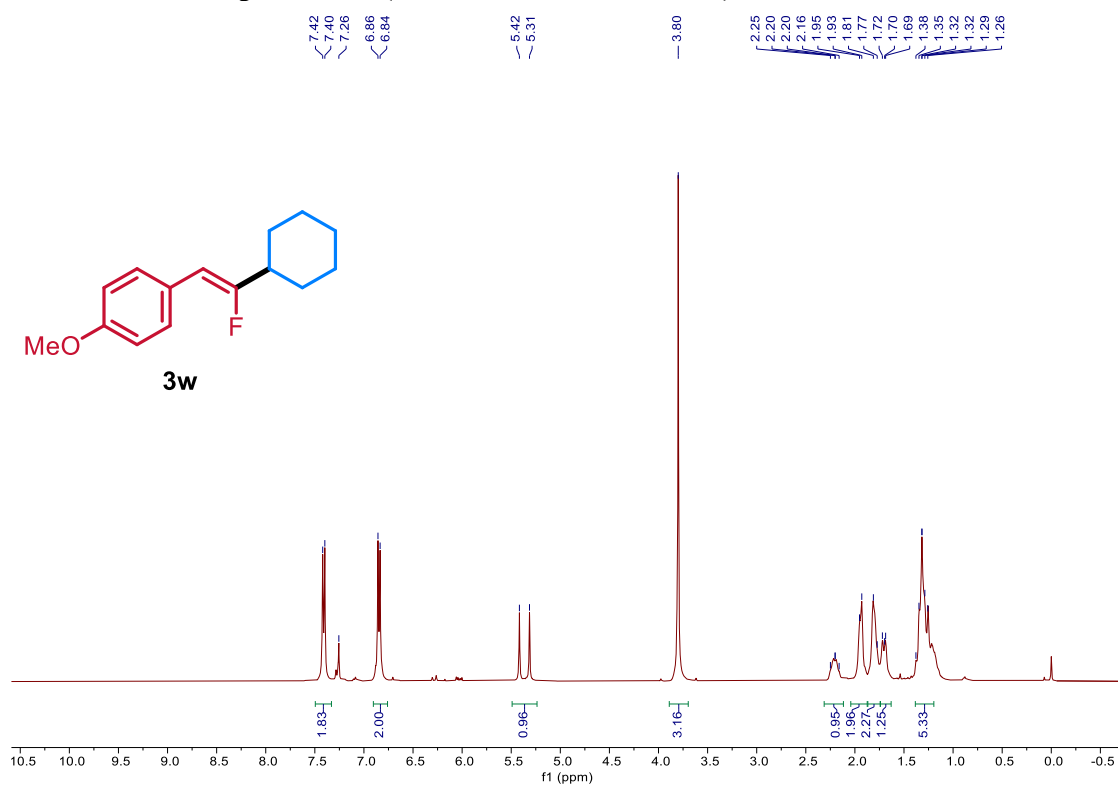
¹³C NMR of Compound 3v (150 MHz, Chloroform-*d*):



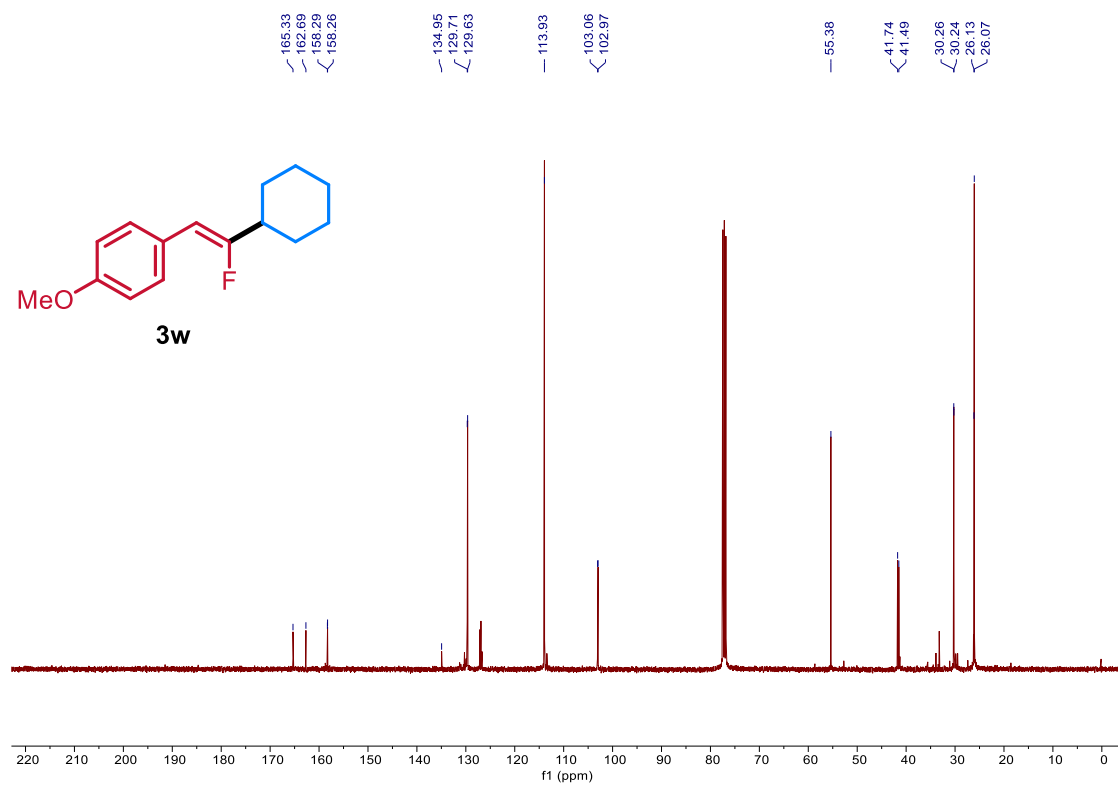
¹⁹F NMR of Compound 3v (375 MHz, Chloroform-*d*):



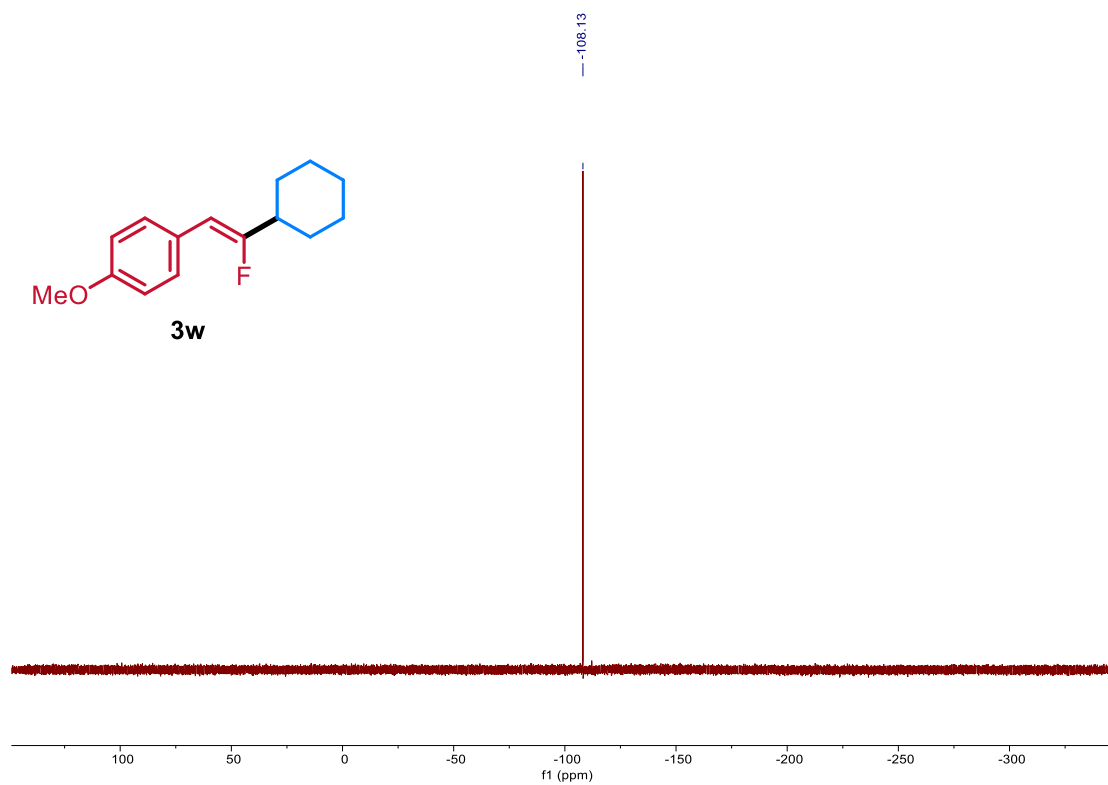
¹H NMR of Compound 3w (400 MHz, Chloroform-*d*):



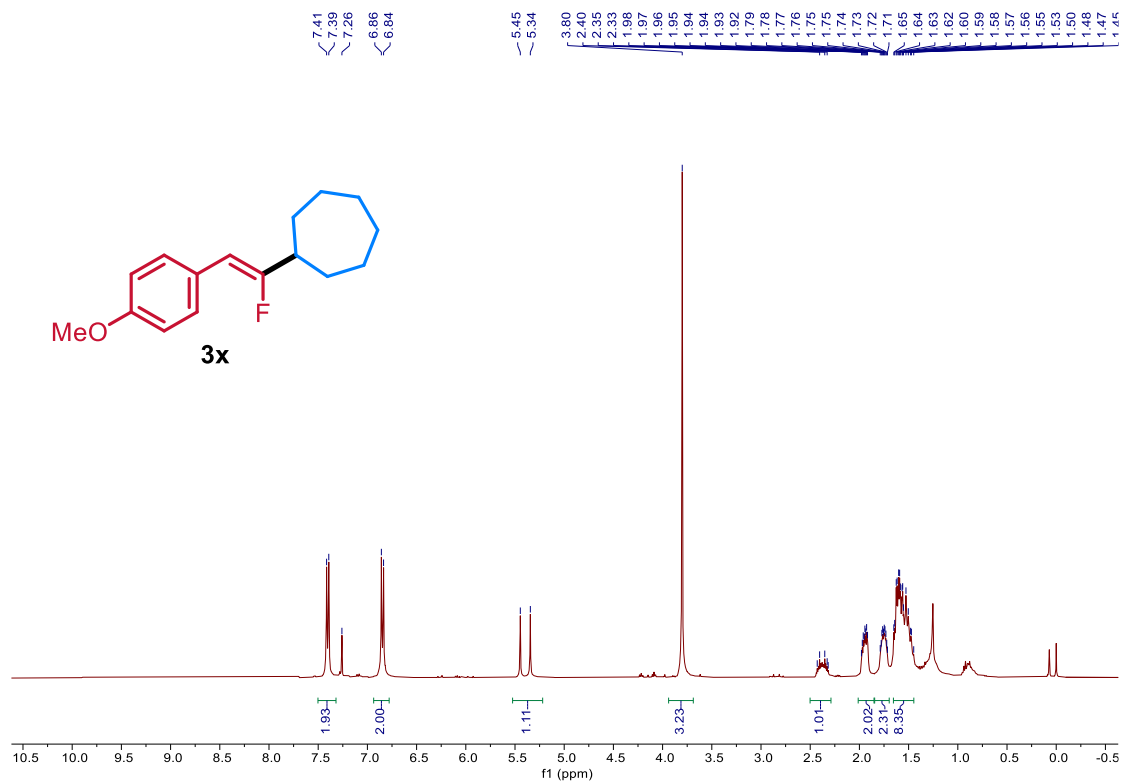
¹³C NMR of Compound 3w (100 MHz, Chloroform-*d*):



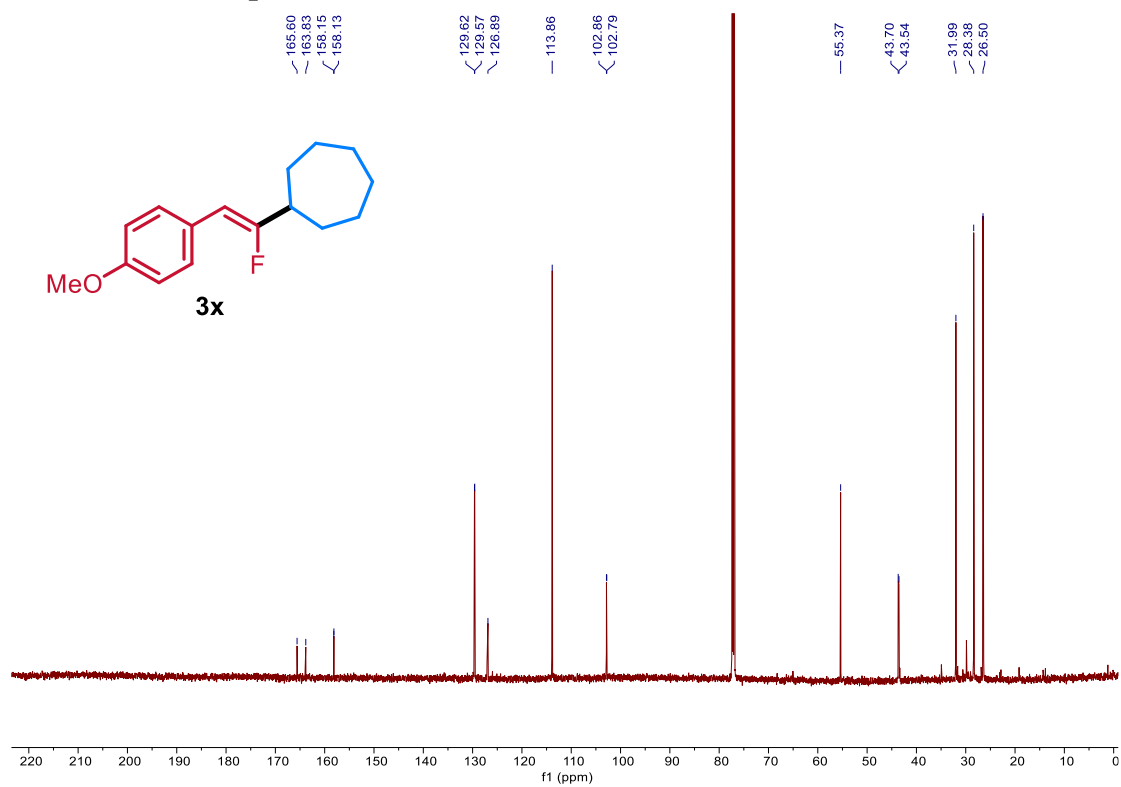
¹⁹F NMR of Compound 3w (375MHz, Chloroform-*d*):



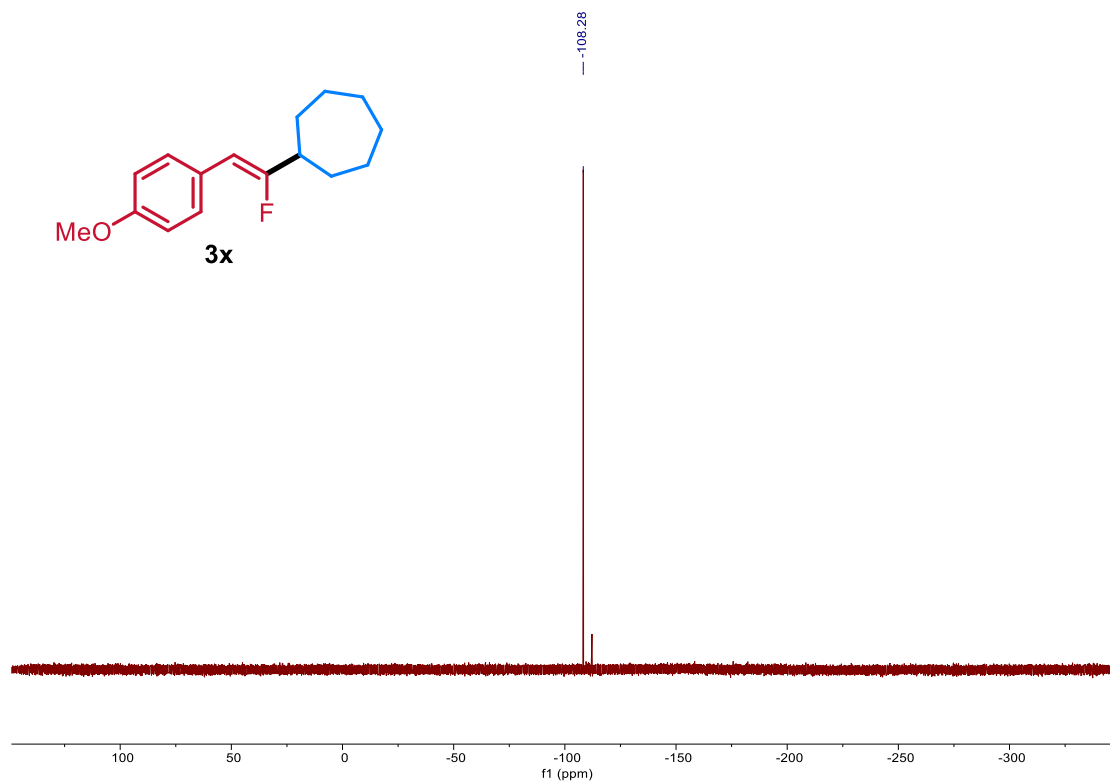
¹H NMR of Compound 3x (400 MHz, Chloroform-*d*):



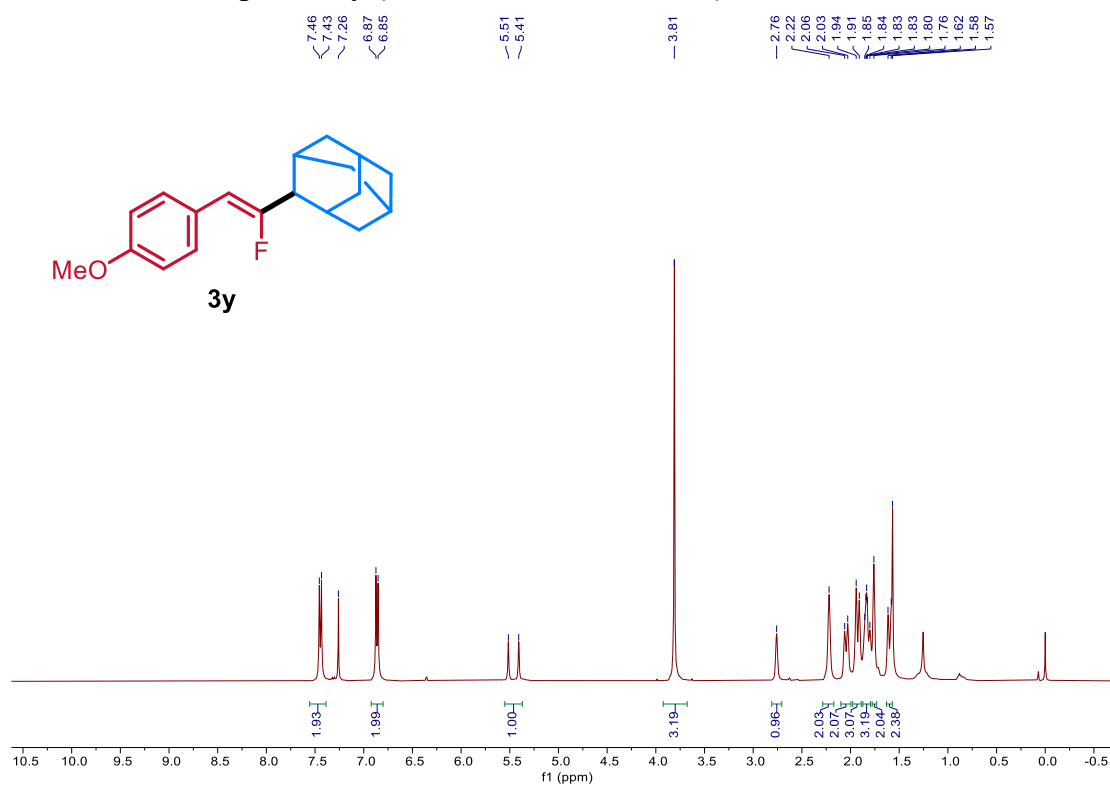
¹³C NMR of Compound 3x (150 MHz, Chloroform-*d*):



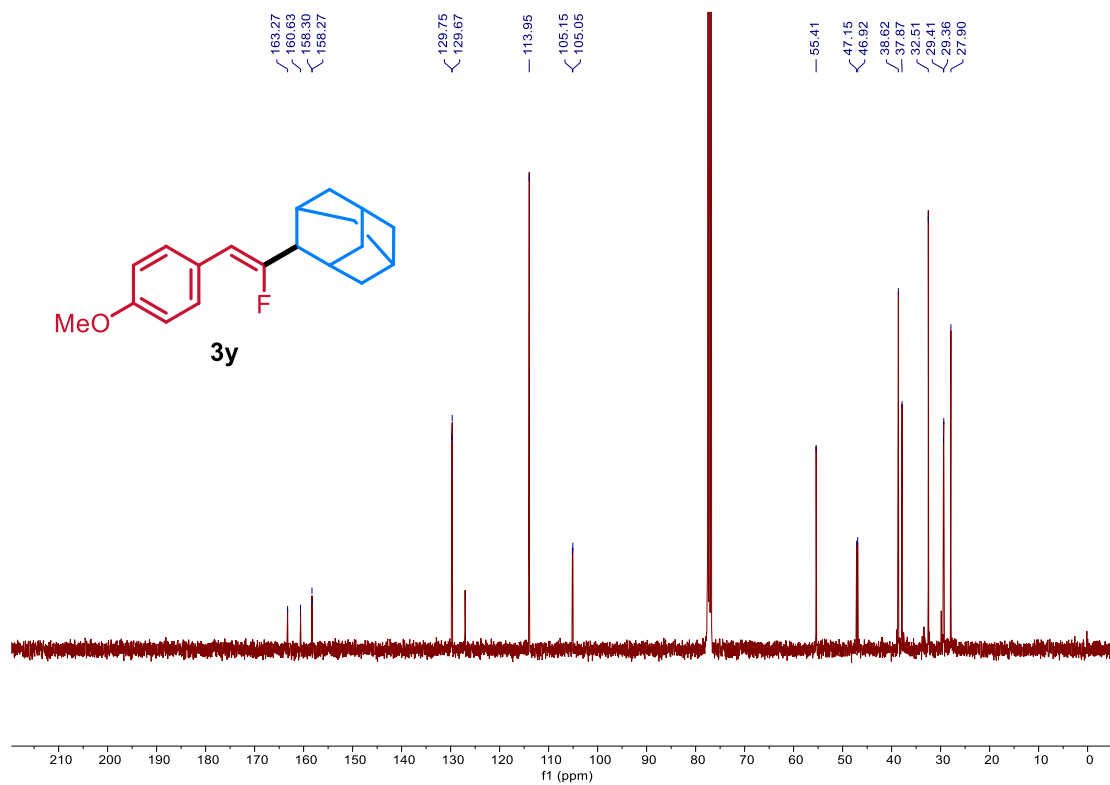
¹⁹F NMR of Compound 3x (375 MHz, Chloroform-*d*):



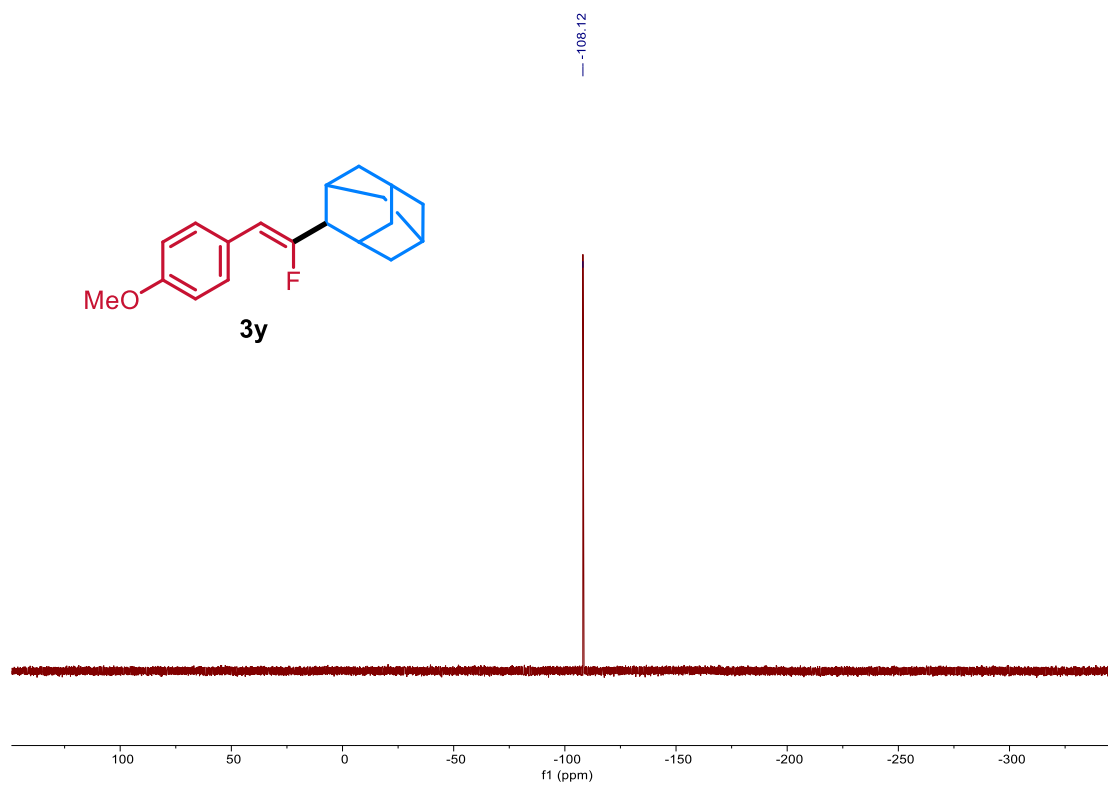
¹H NMR of Compound 3y (400 MHz, Chloroform-*d*):



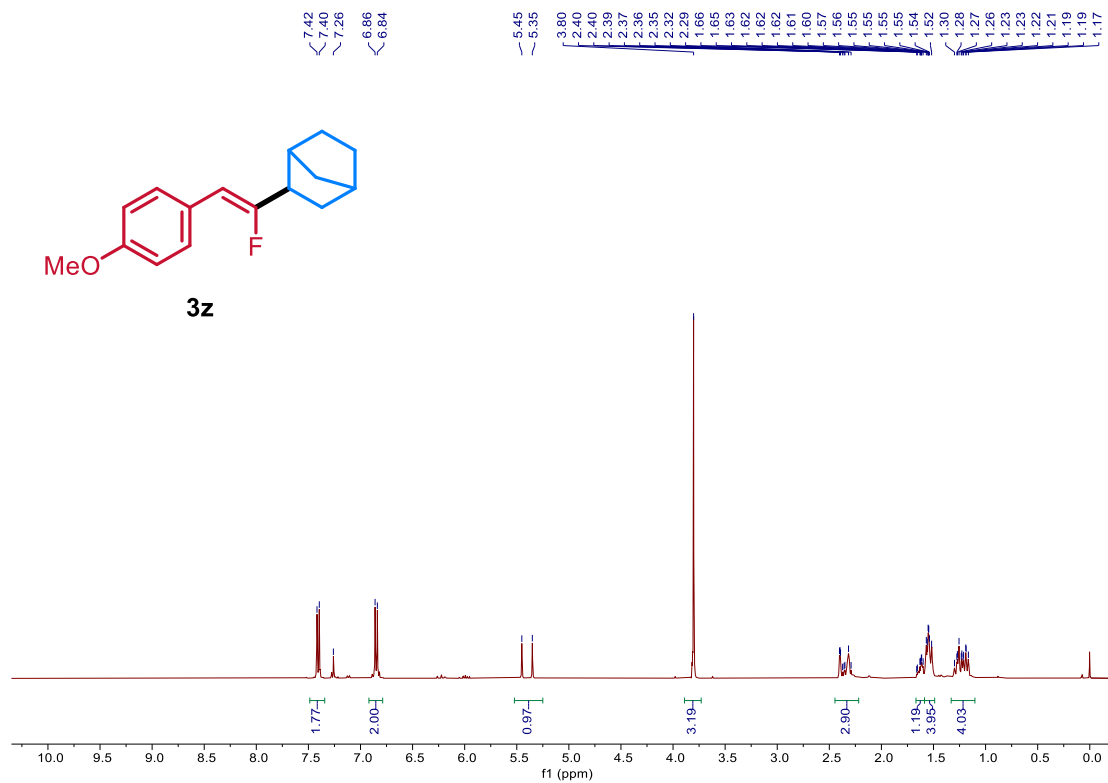
¹³C NMR of Compound 3y (100 MHz, Chloroform-*d*):



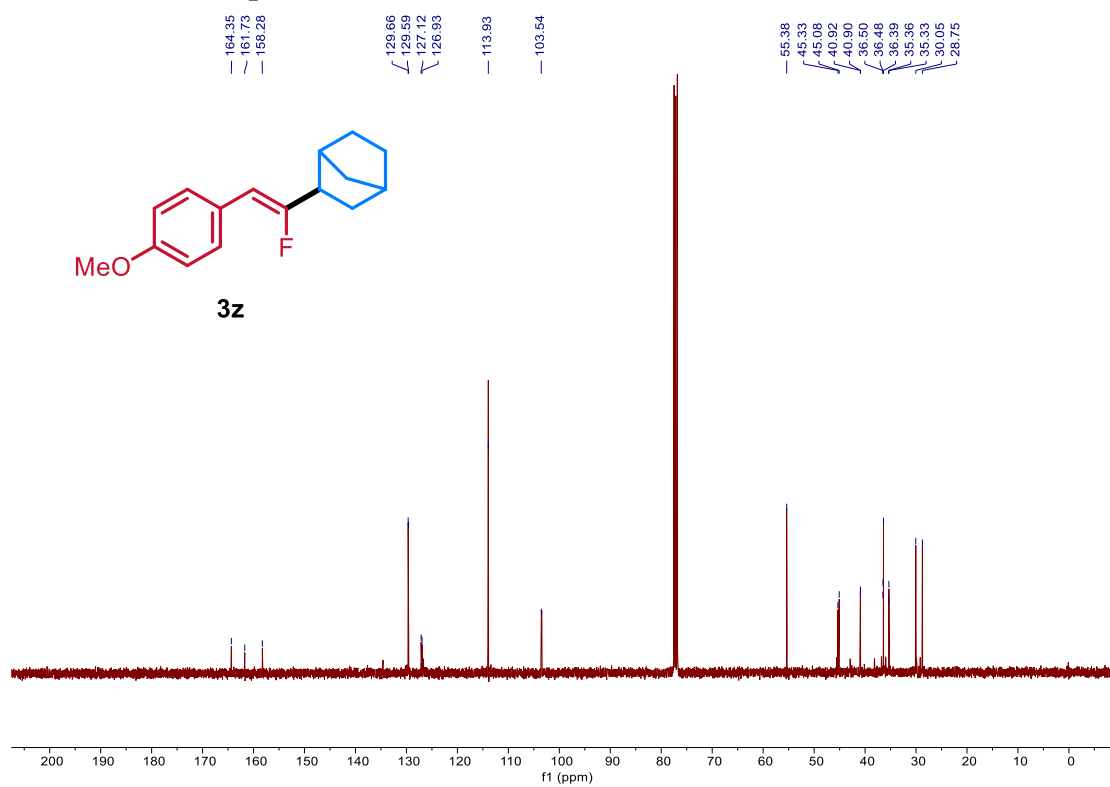
¹⁹F NMR of Compound 3y (375 MHz, Chloroform-*d*):



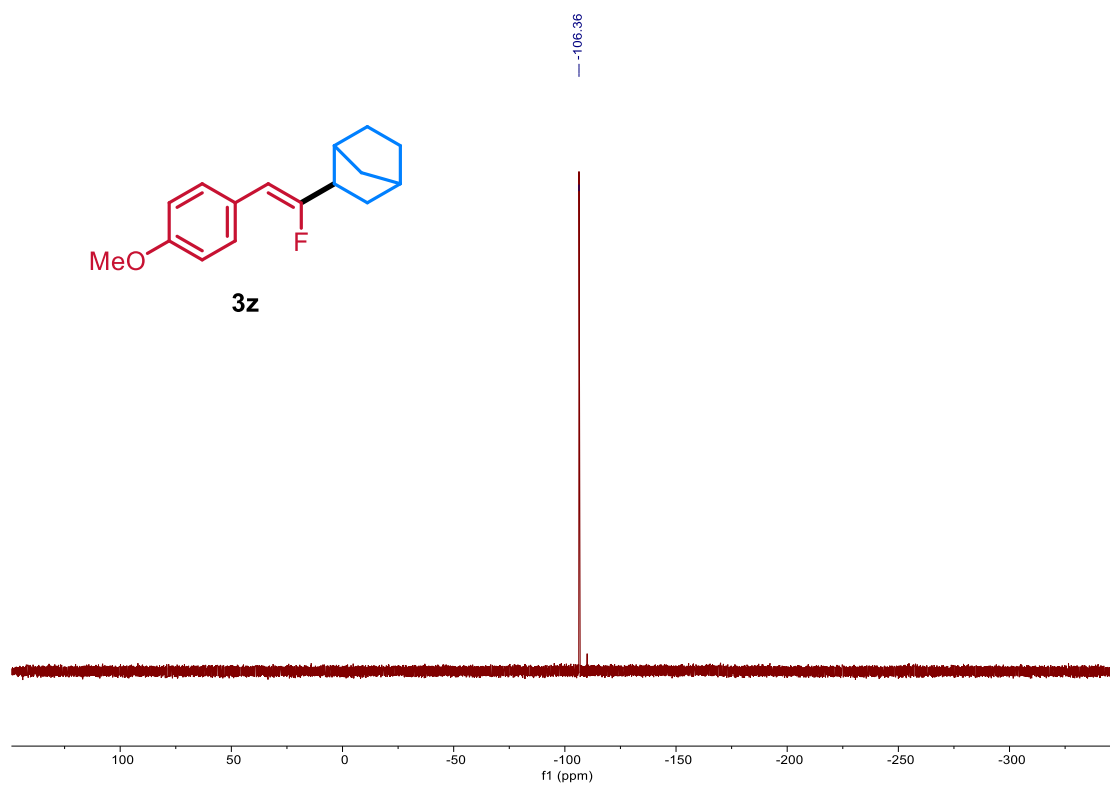
¹H NMR of Compound 3z (400 MHz, Chloroform-*d*):



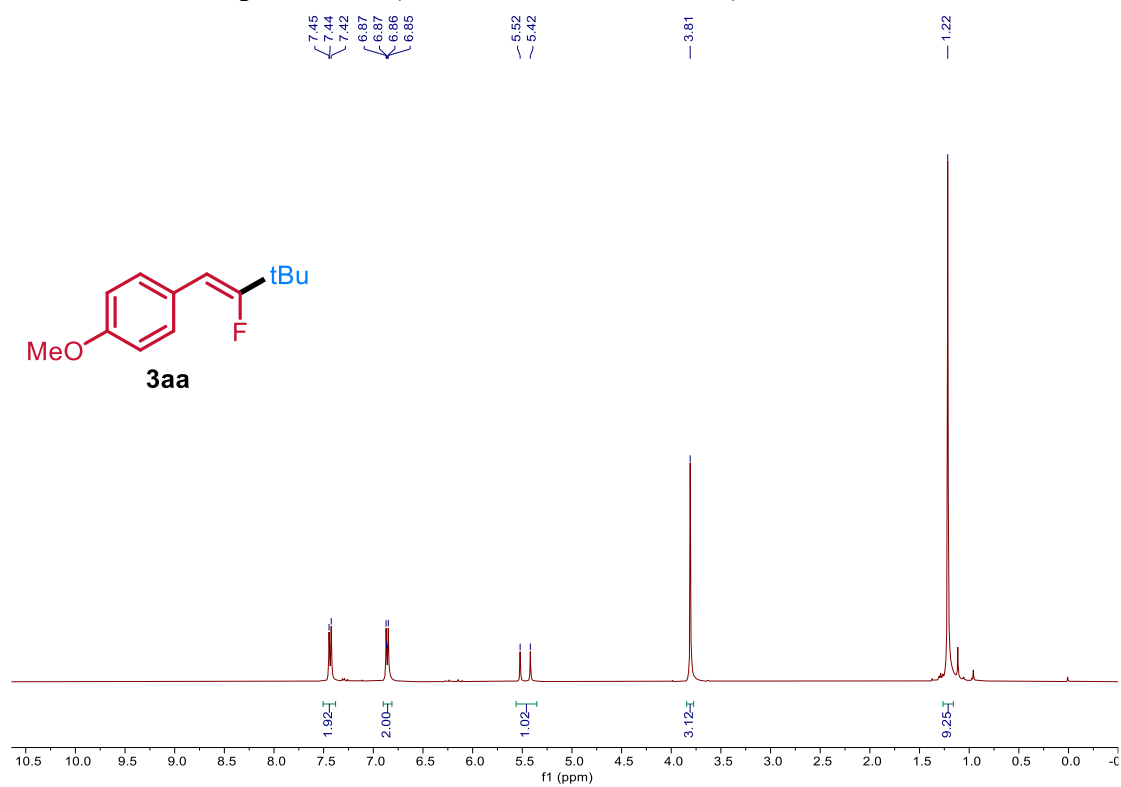
¹³C NMR of Compound 3z (100 MHz, Chloroform-*d*):



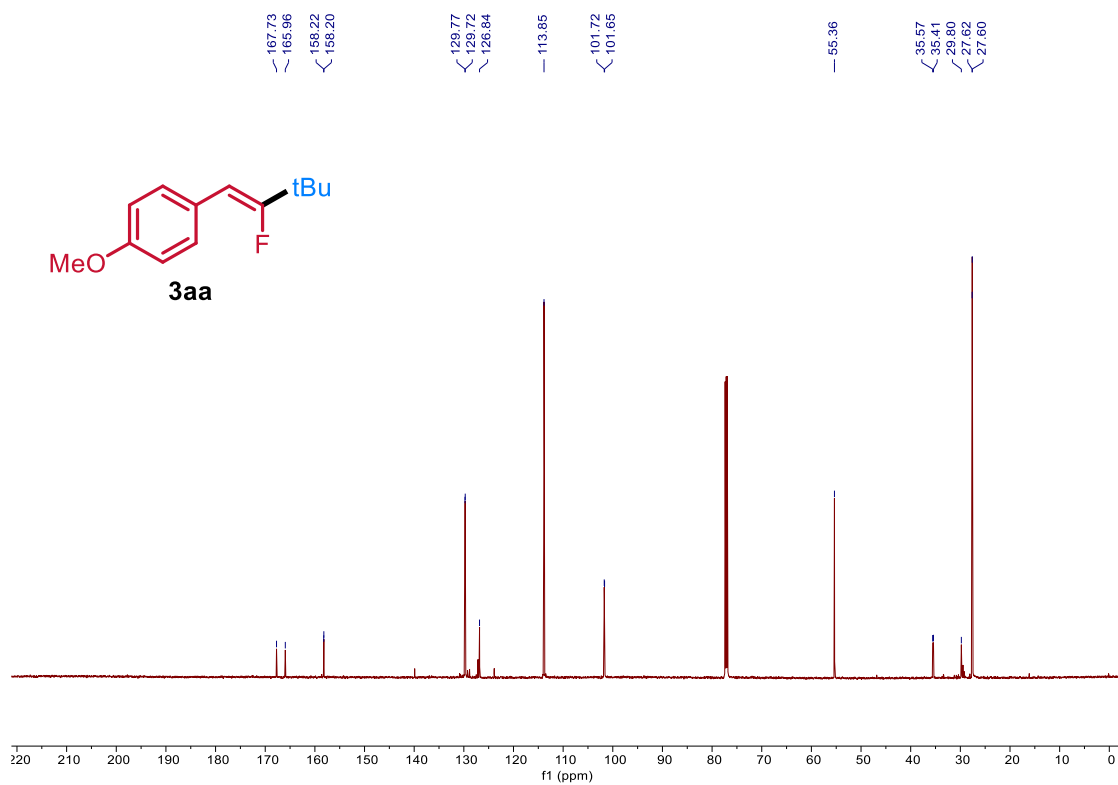
¹⁹F NMR of Compound 3z (375 MHz, Chloroform-*d*):



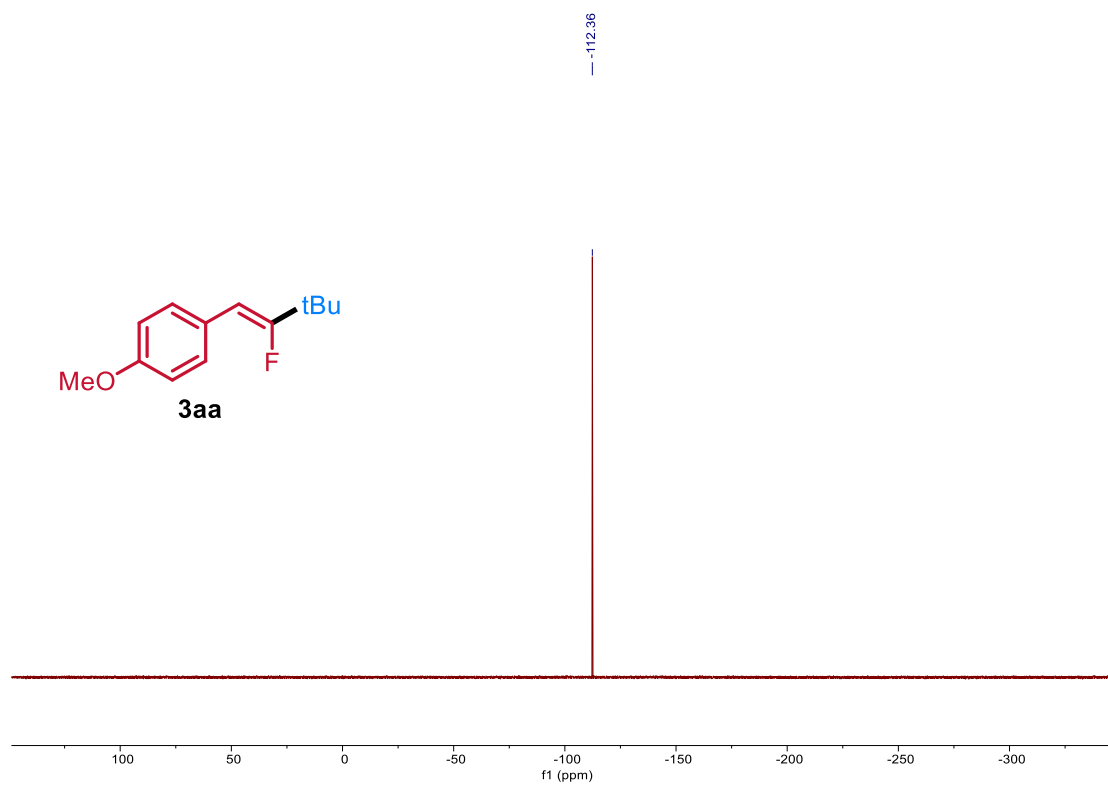
¹H NMR of Compound 3aa (400 MHz, Chloroform-*d*):



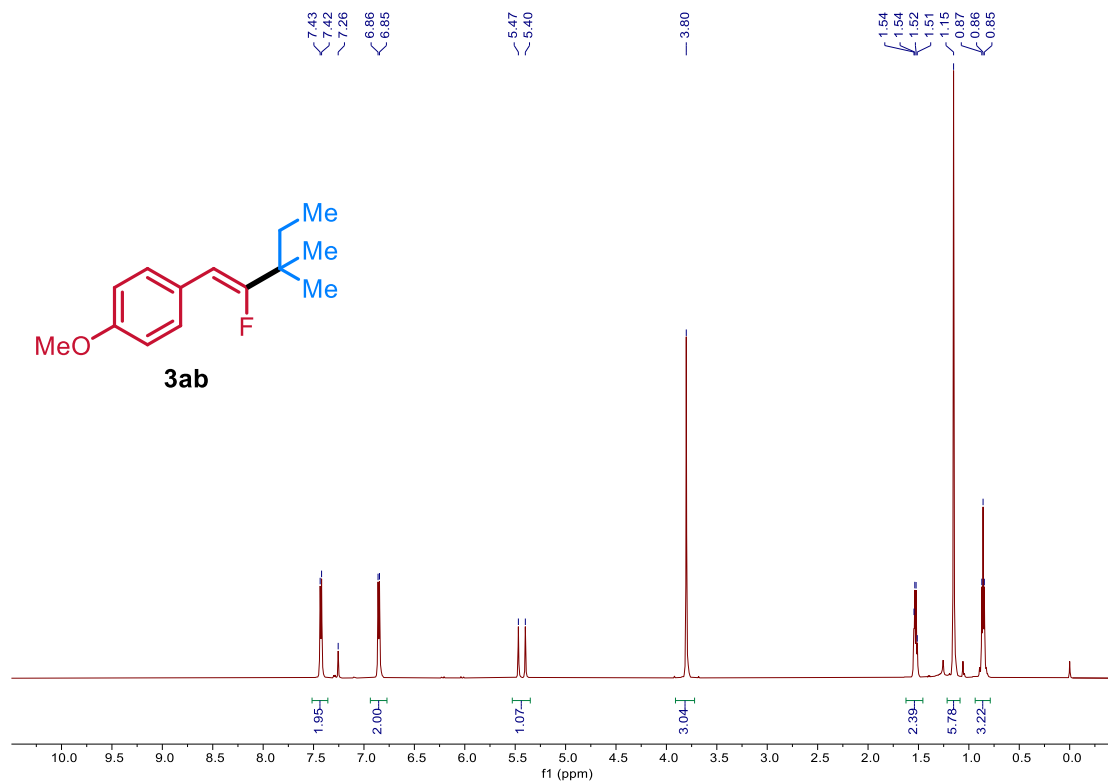
¹³C NMR of Compound 3aa (100 MHz, Chloroform-*d*):



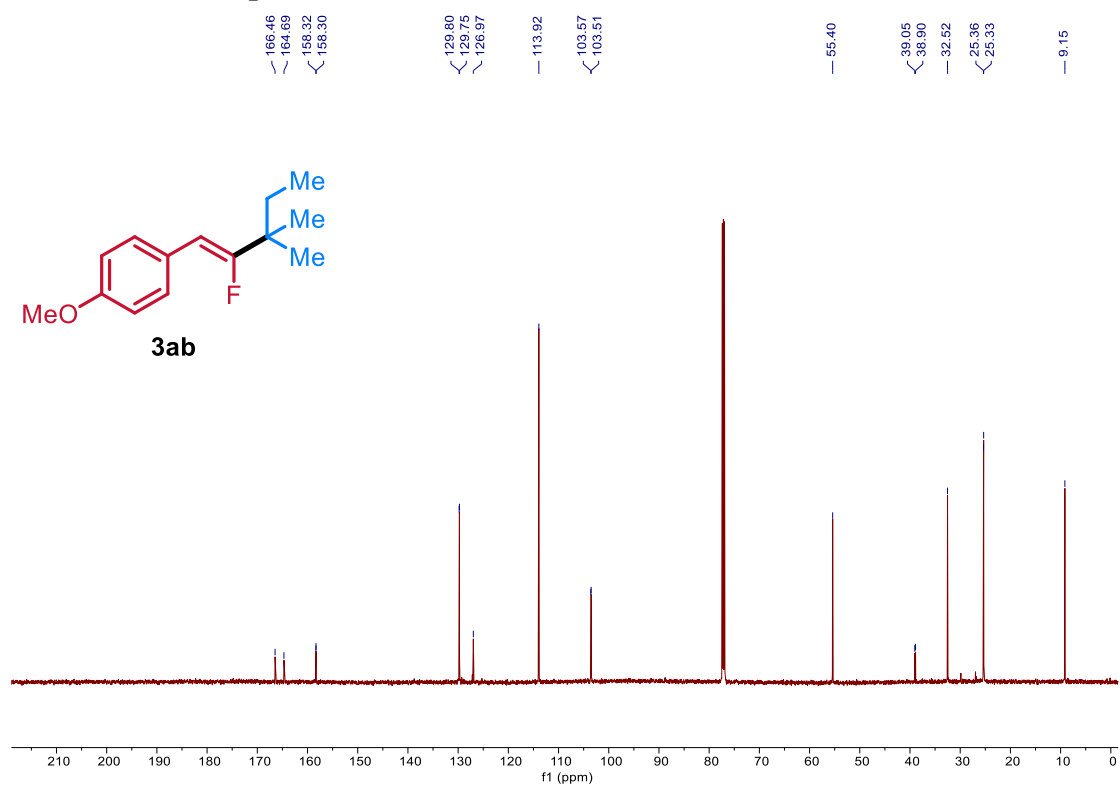
¹⁹F NMR of Compound 3aa (375 MHz, Chloroform-*d*):



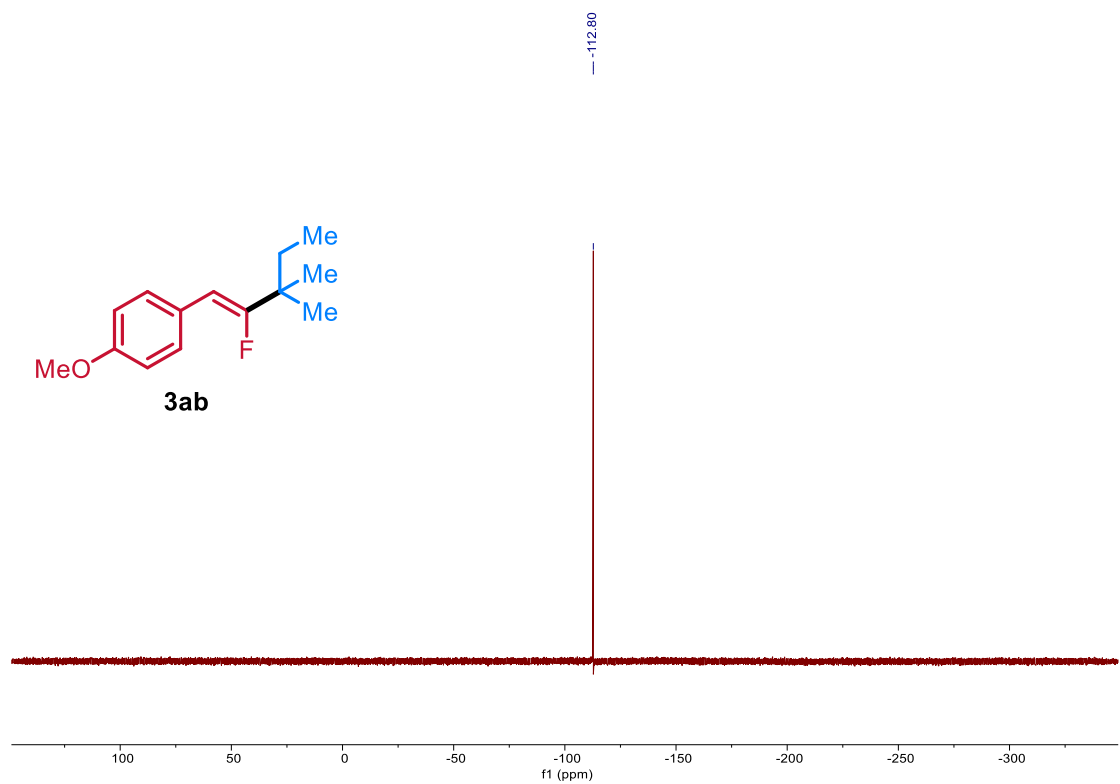
¹H NMR of Compound 3ab (600 MHz, Chloroform-*d*):



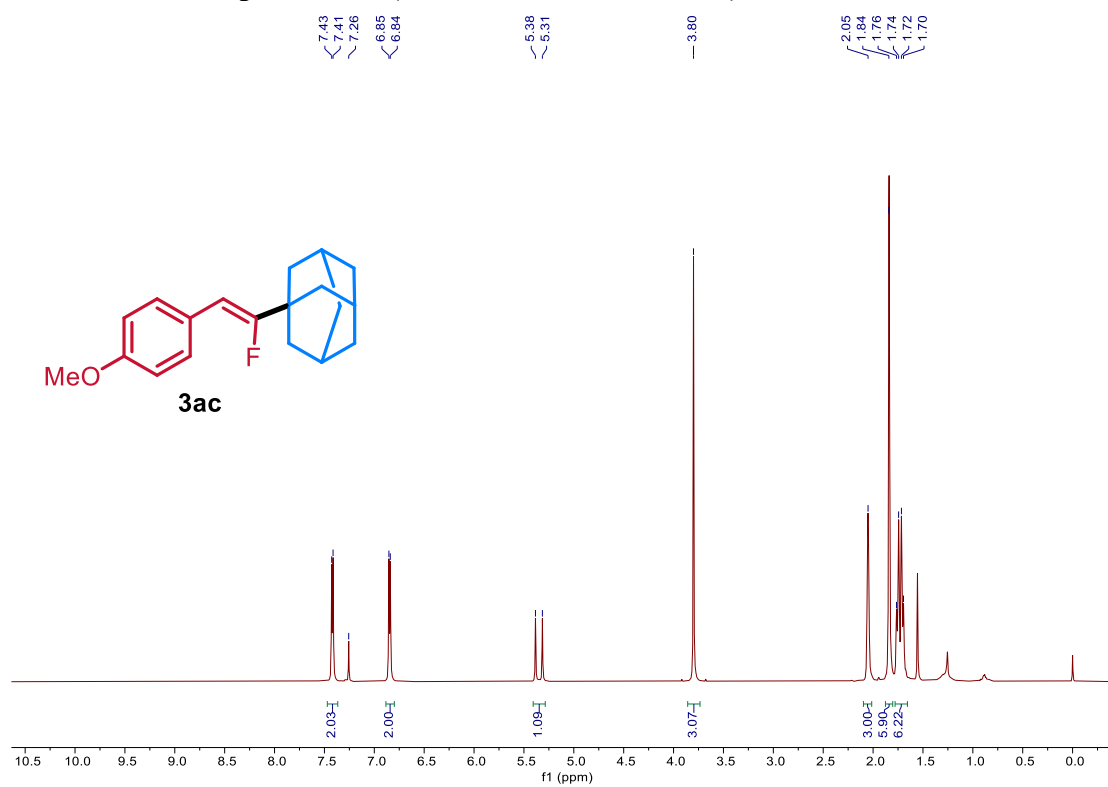
¹³C NMR of Compound 3ab (150 MHz, Chloroform-*d*):



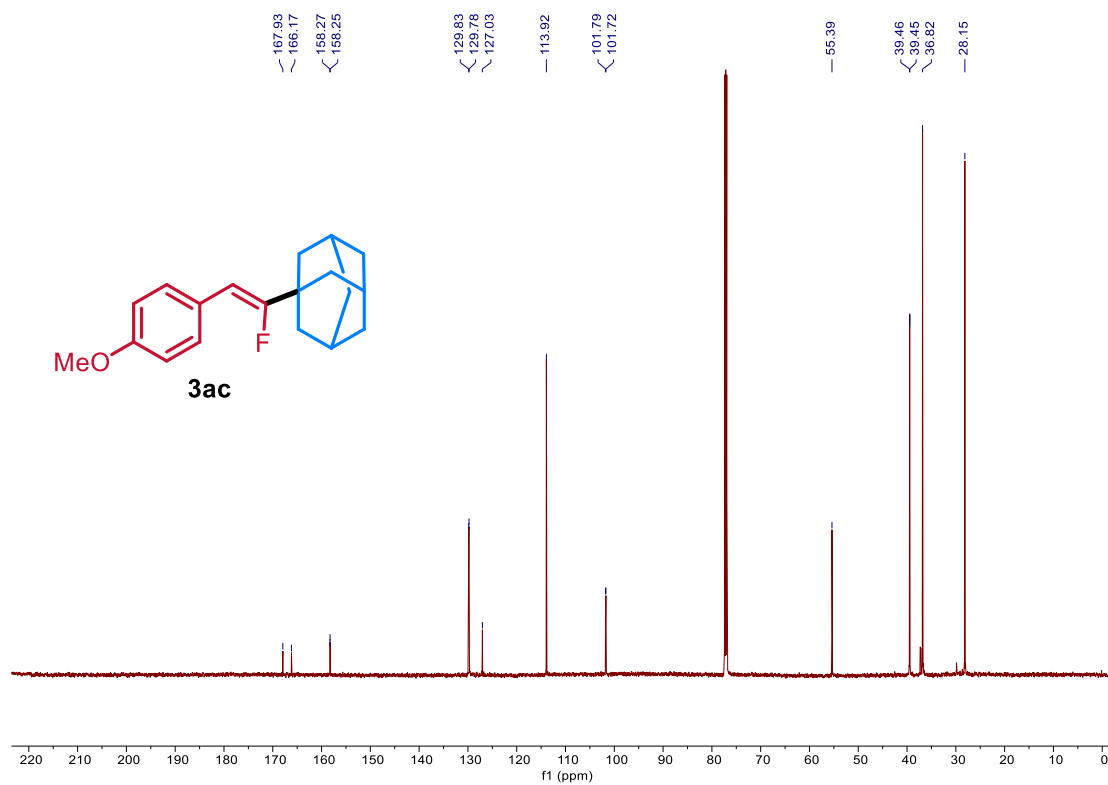
¹⁹F NMR of Compound 3ab (375 MHz, Chloroform-*d*):



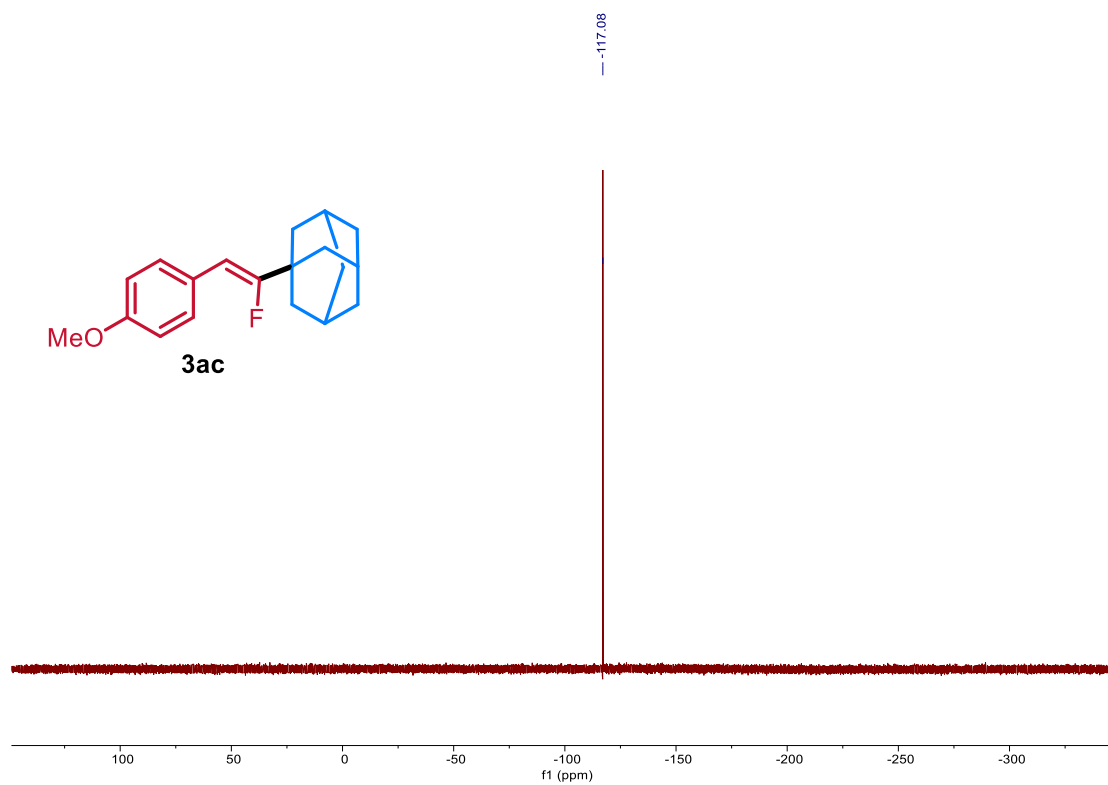
¹H NMR of Compound 3ac (600 MHz, Chloroform-*d*):



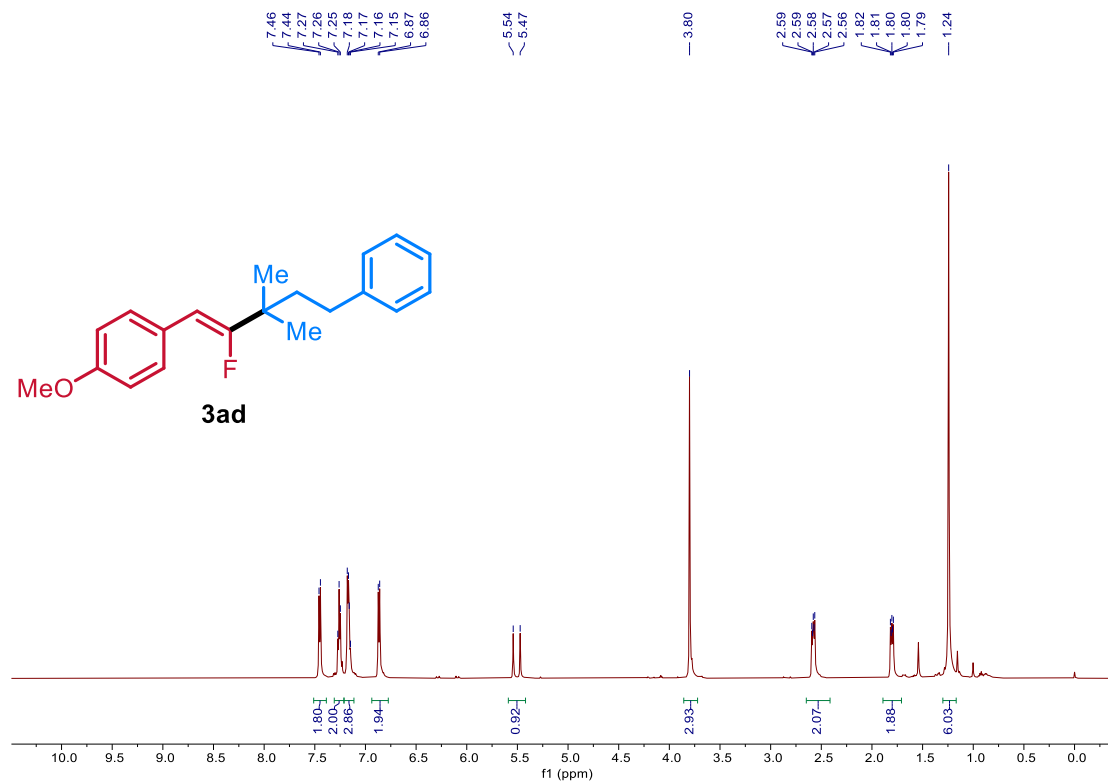
¹³C NMR of Compound 3ac (150 MHz, Chloroform-*d*):



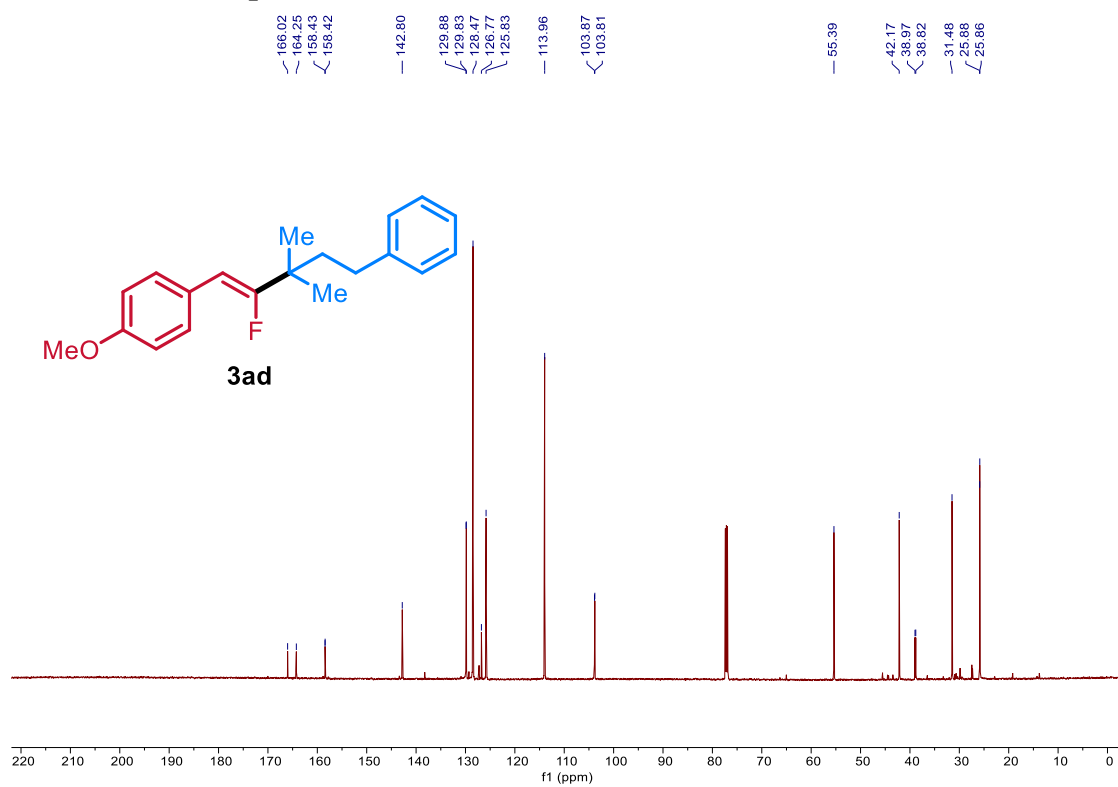
¹⁹F NMR of Compound 3ac (375 MHz, Chloroform-*d*):



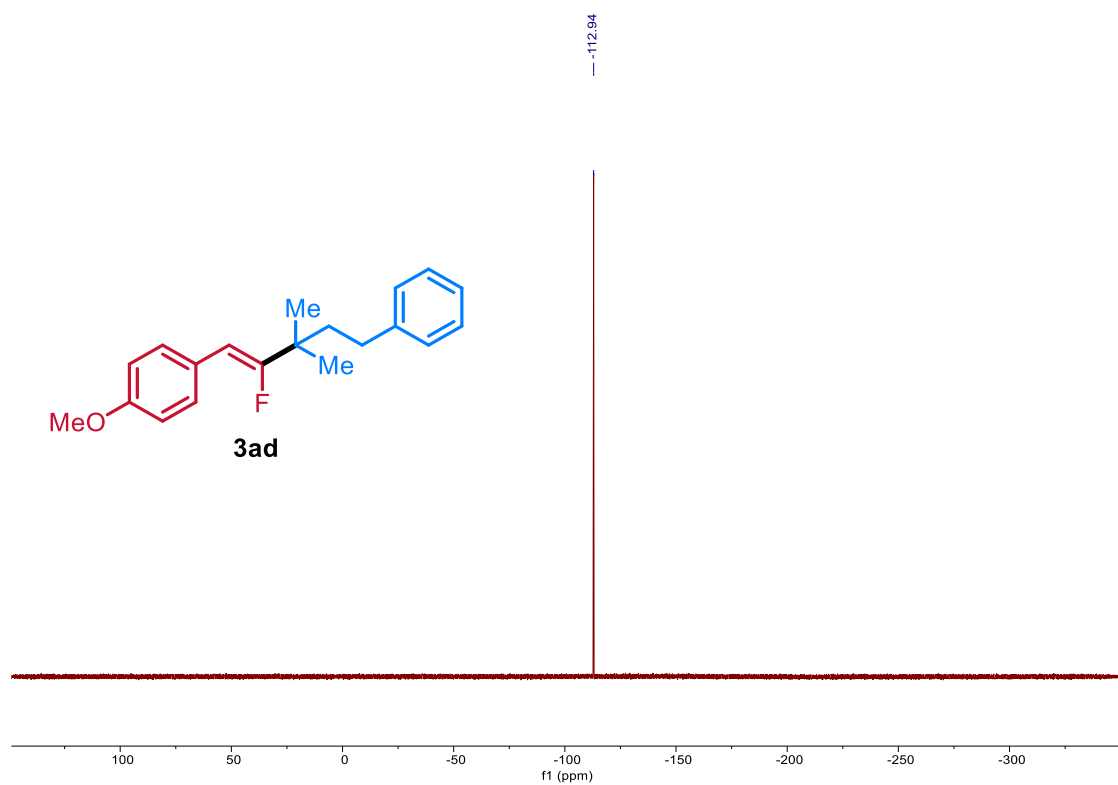
¹H NMR of Compound 3ad (600 MHz, Chloroform-*d*):



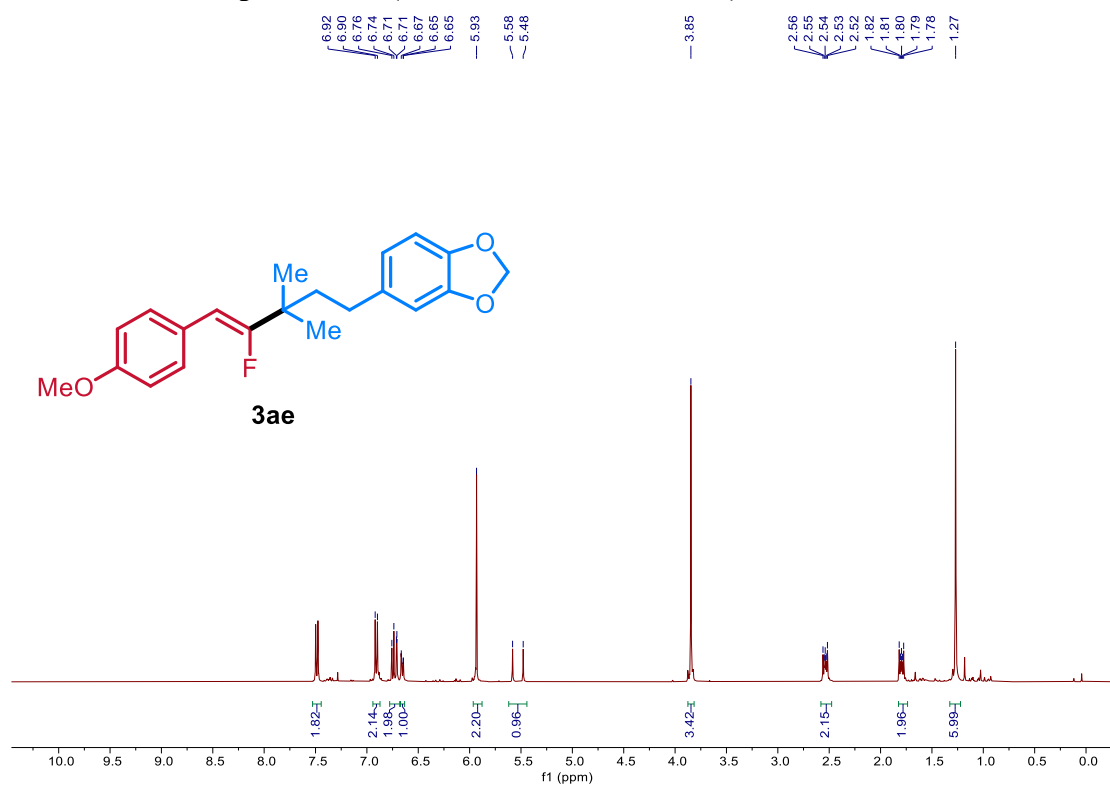
¹³C NMR of Compound 3ad (150 MHz, Chloroform-*d*):



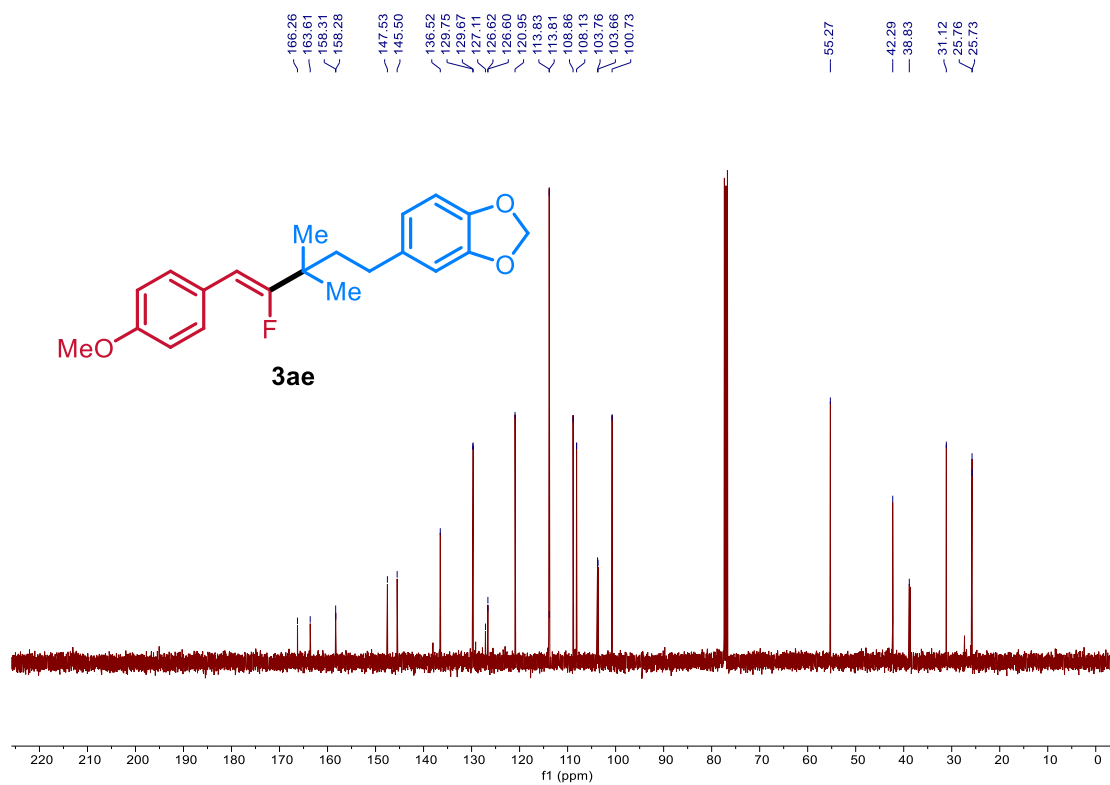
¹⁹F NMR of Compound 3ad (375 MHz, Chloroform-*d*):



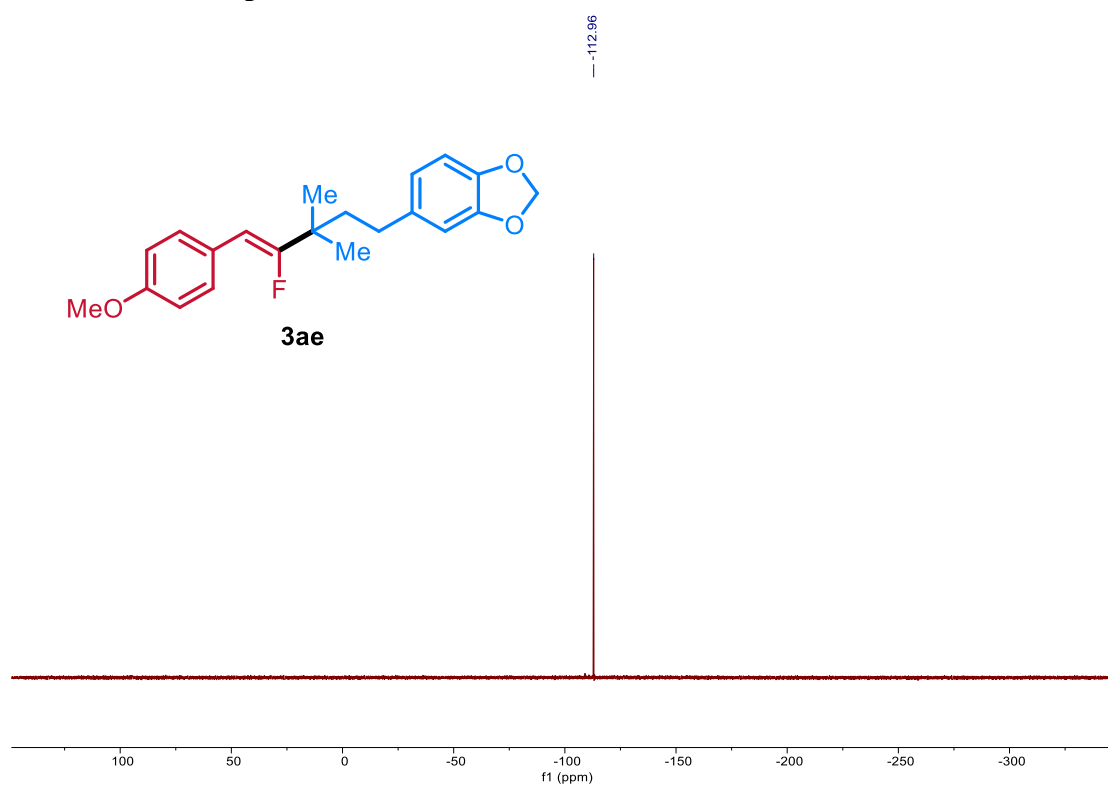
¹H NMR of Compound 3ae (600 MHz, Chloroform-*d*):



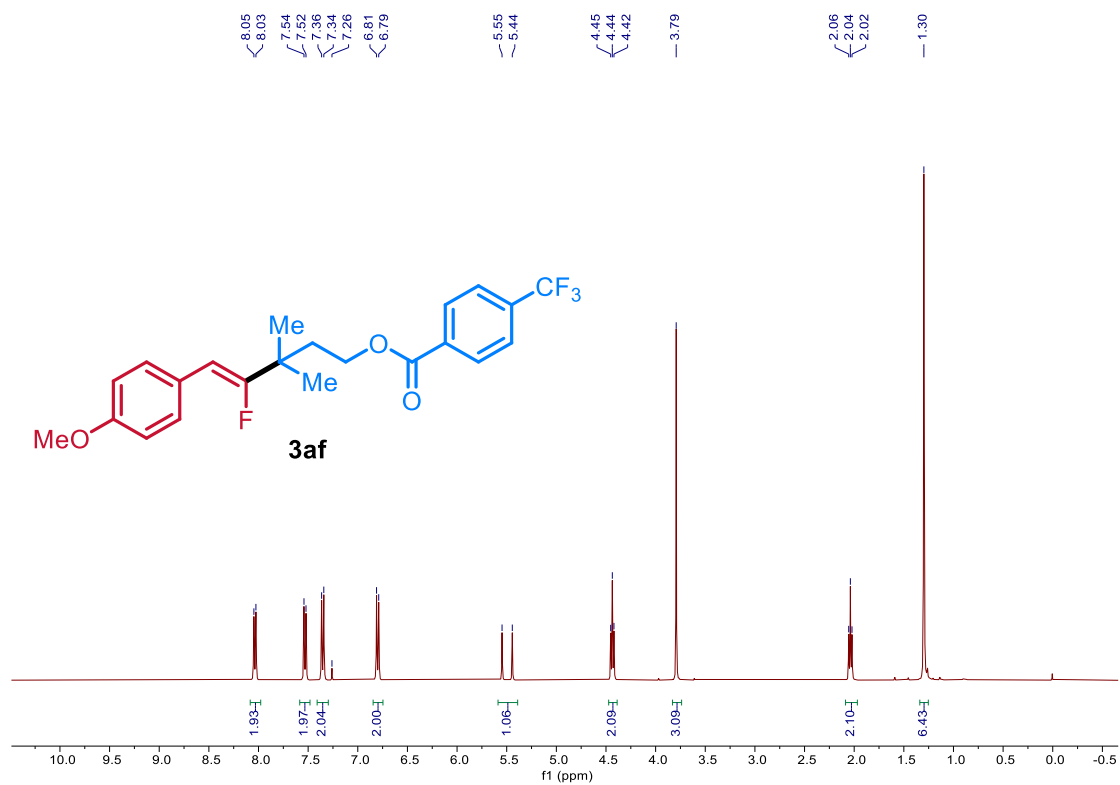
¹³C NMR of Compound 3ae (150 MHz, Chloroform-*d*):



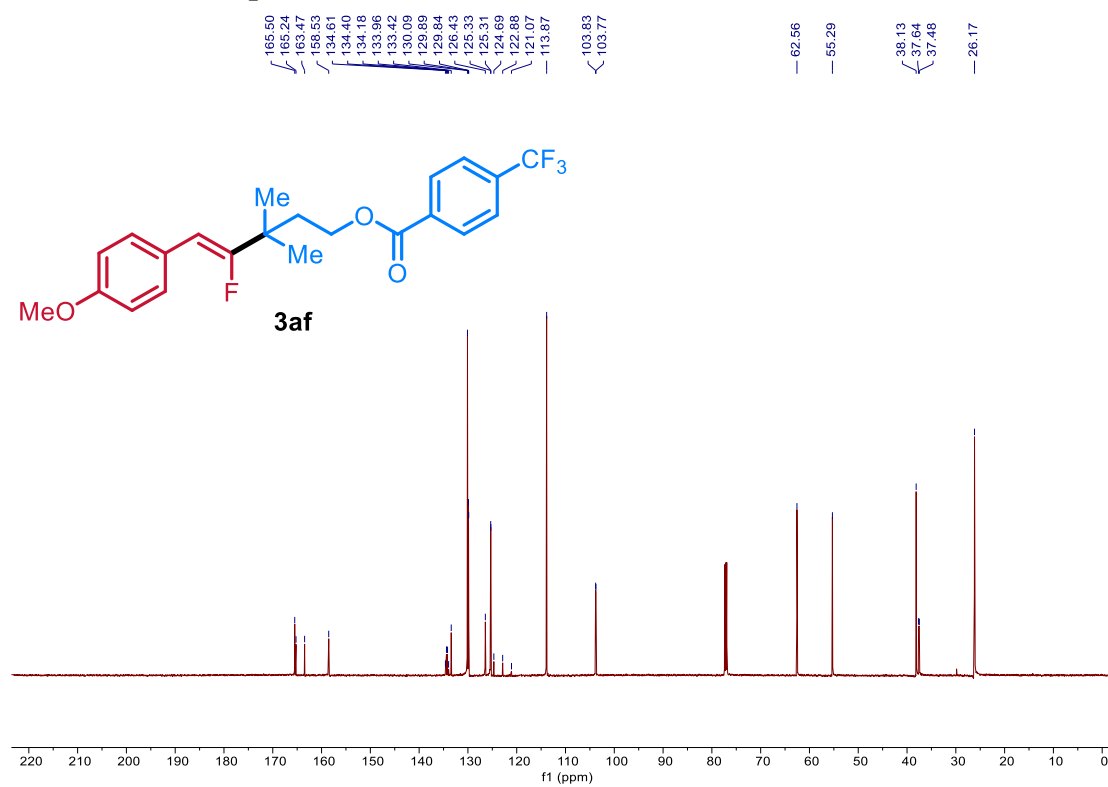
¹⁹F NMR of Compound 3ae (375 MHz, Chloroform-*d*):



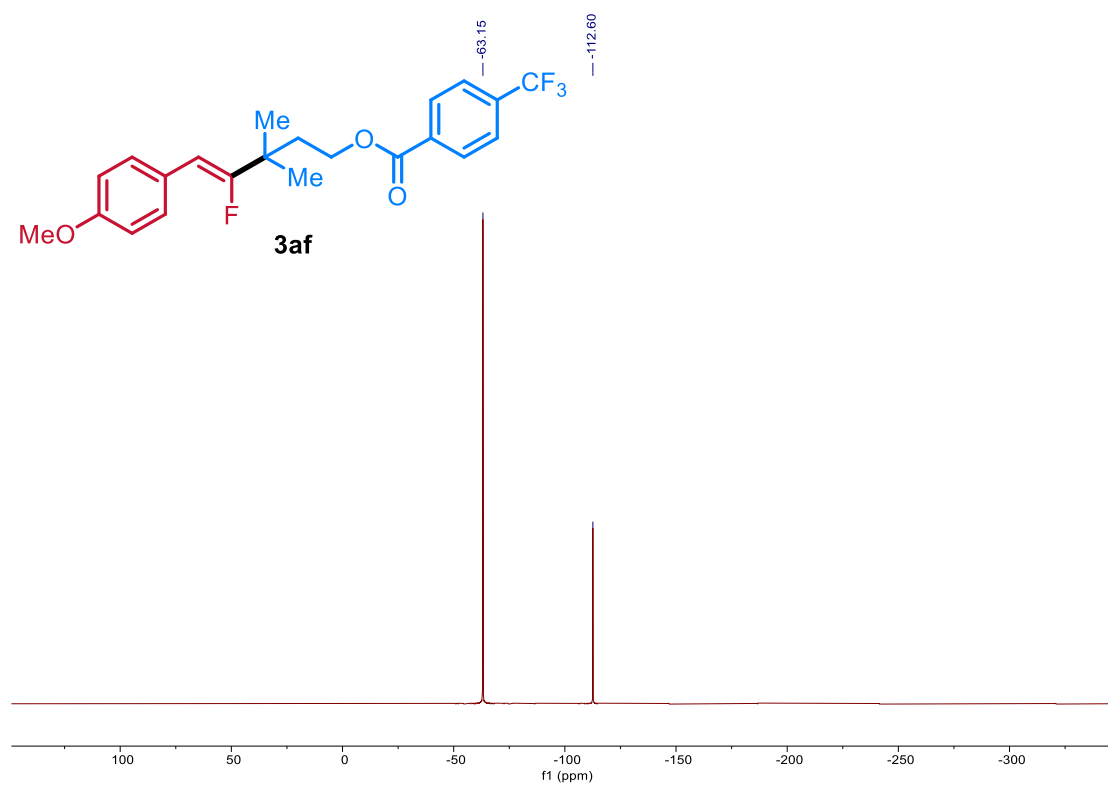
¹H NMR of Compound 3af (400 MHz, Chloroform-*d*):



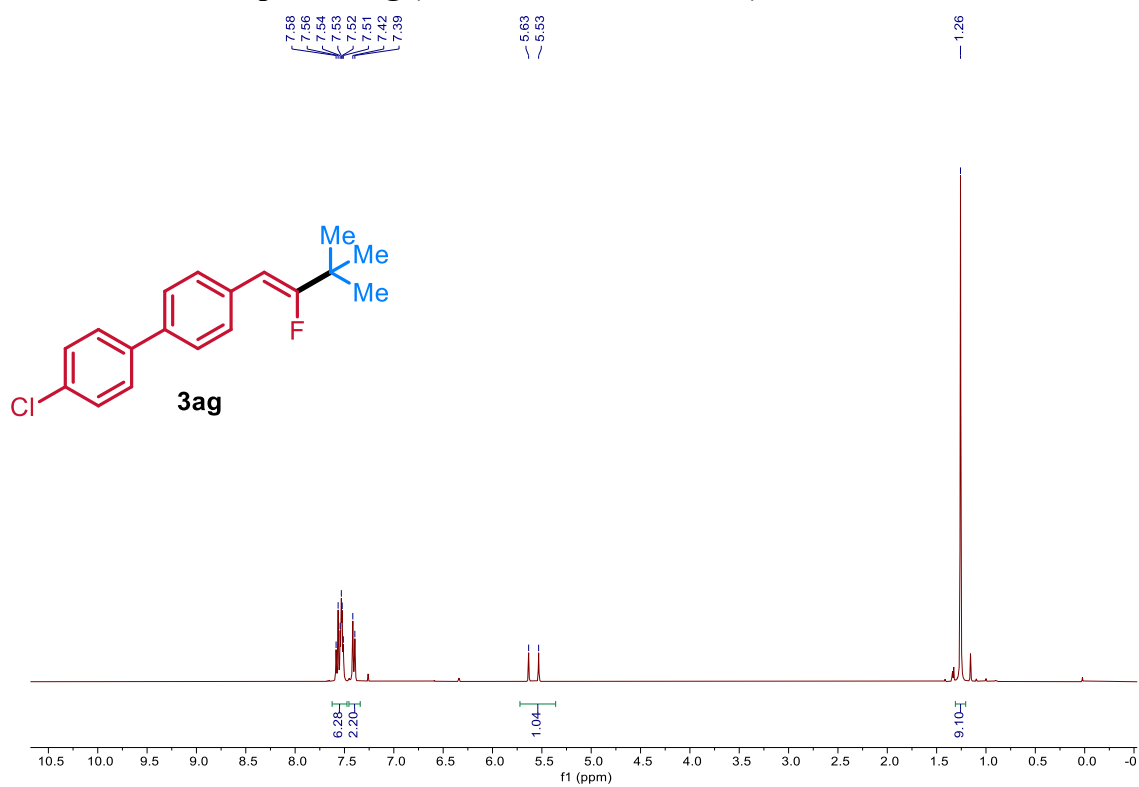
¹³C NMR of Compound 3ad (150 MHz, Chloroform-*d*):



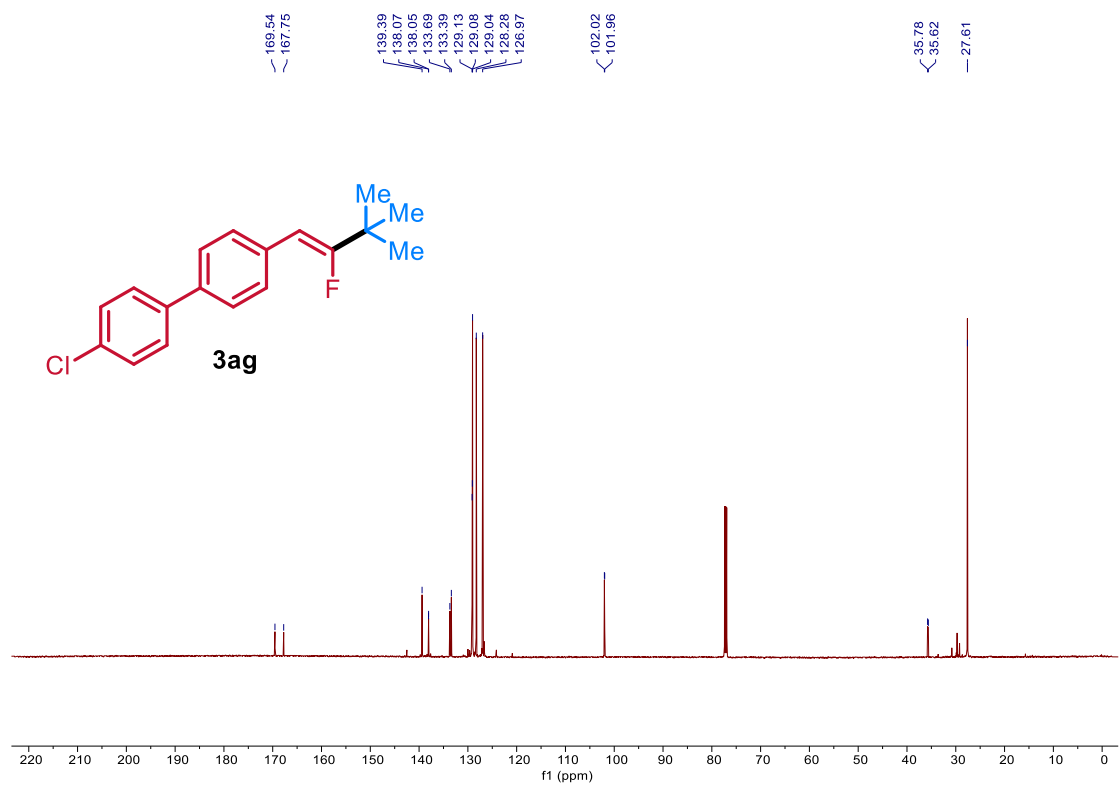
¹⁹F NMR of Compound 3ad (375 MHz, Chloroform-*d*):



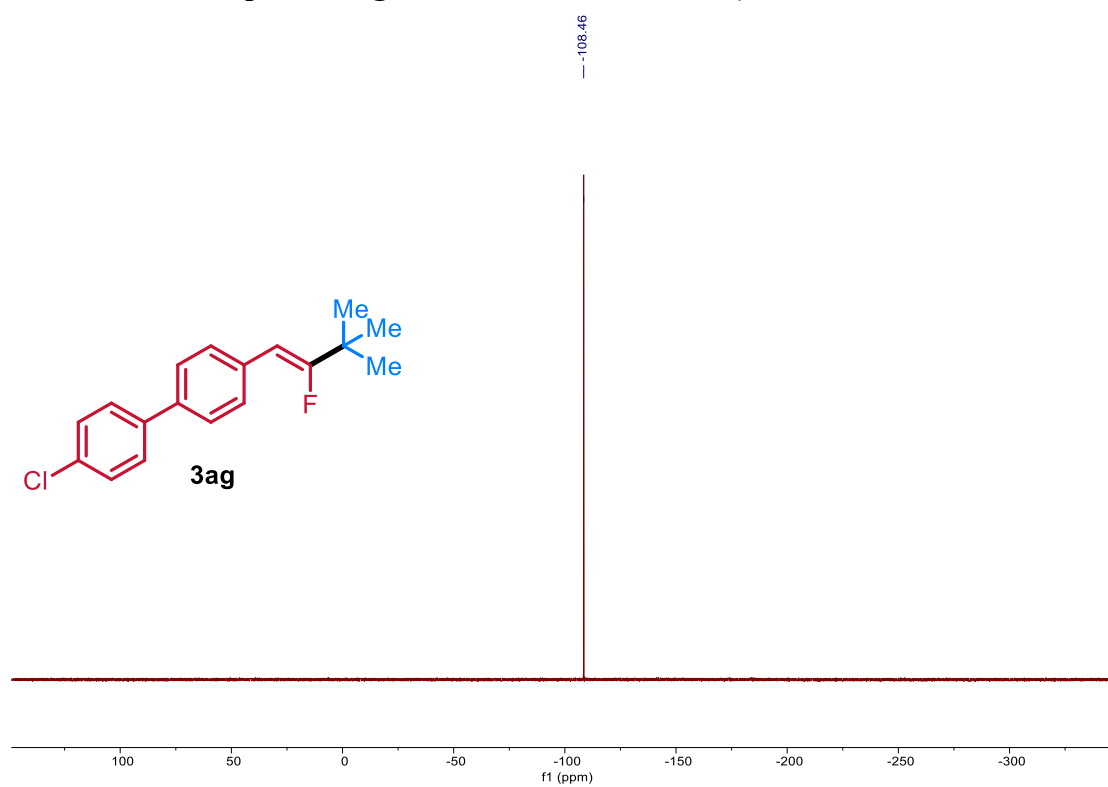
¹H NMR of Compound 3ag (400 MHz, Chloroform-*d*):



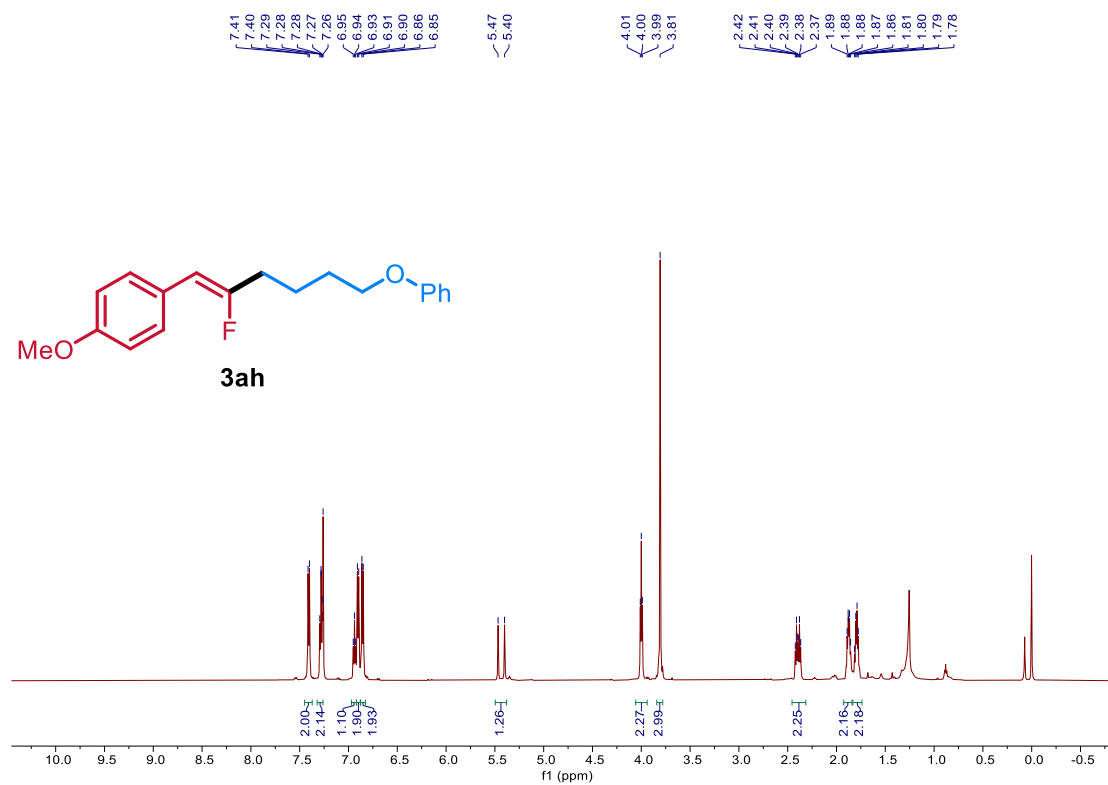
¹³C NMR of Compound 3ag (150 MHz, Chloroform-*d*):



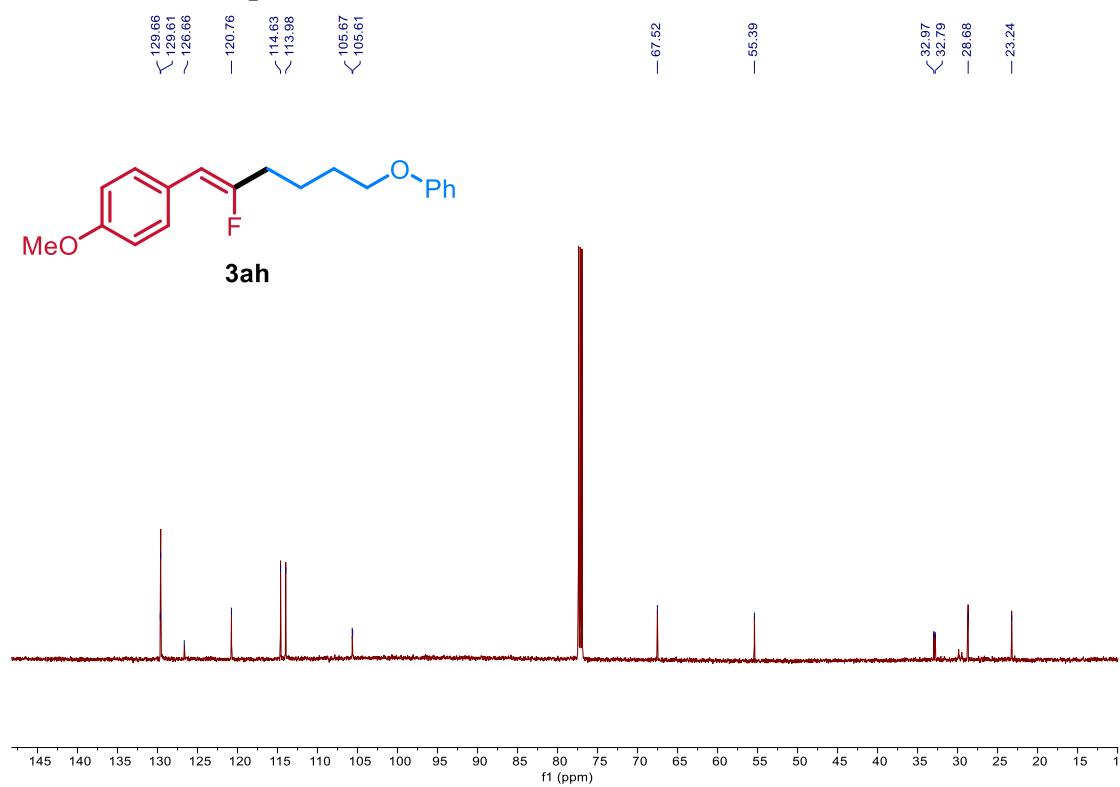
^{19}F NMR of Compound 3ag (375 MHz, Chloroform-*d*): δ - 108.46 (s, 1F)



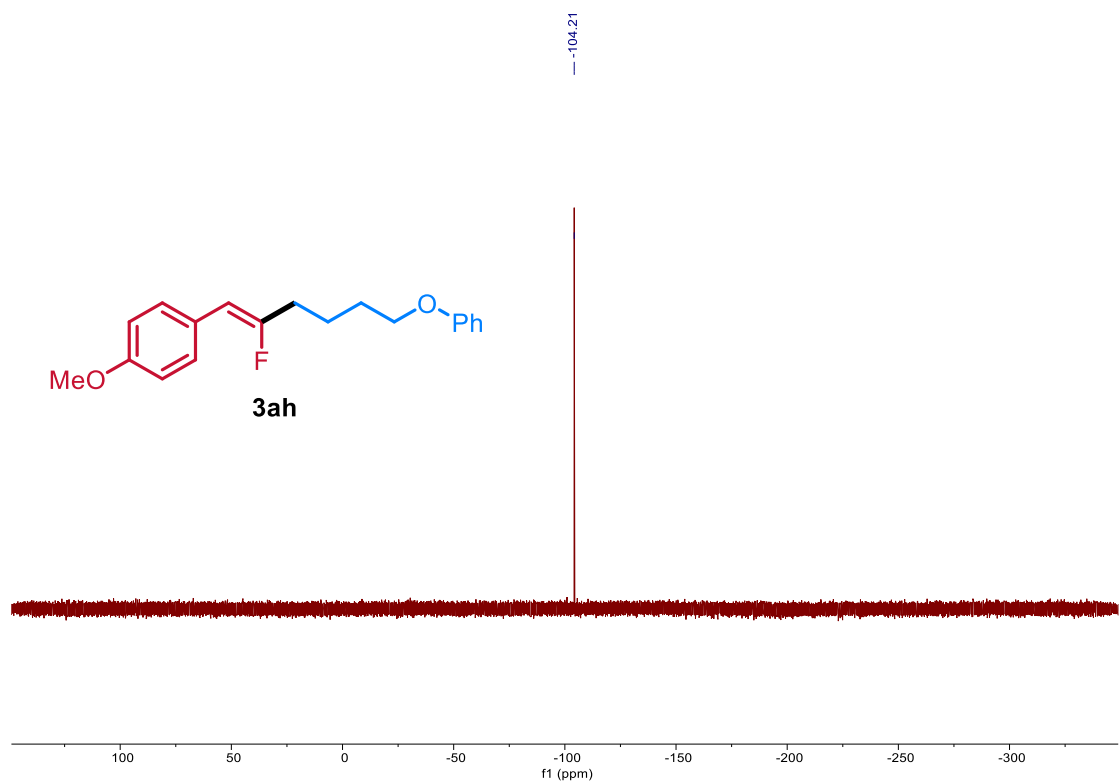
^1H NMR of Compound 3ah (400 MHz, Chloroform-*d*):



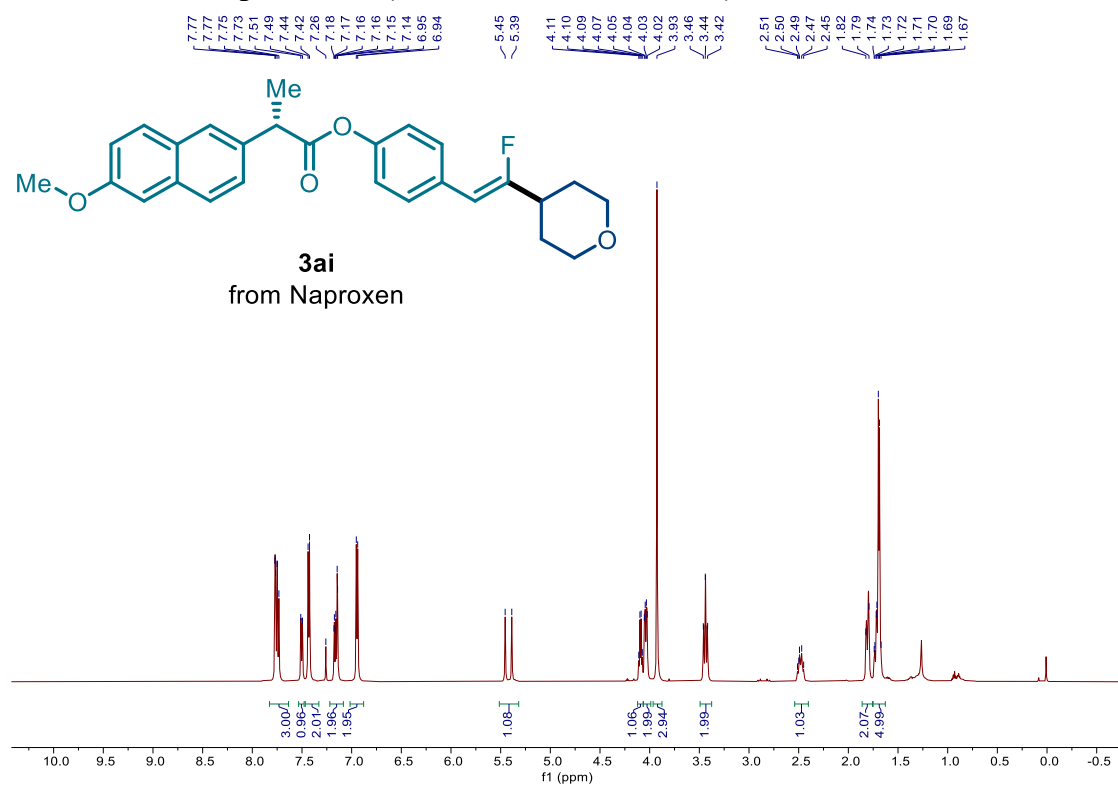
¹³C NMR of Compound 3ah (150 MHz, Chloroform-*d*):



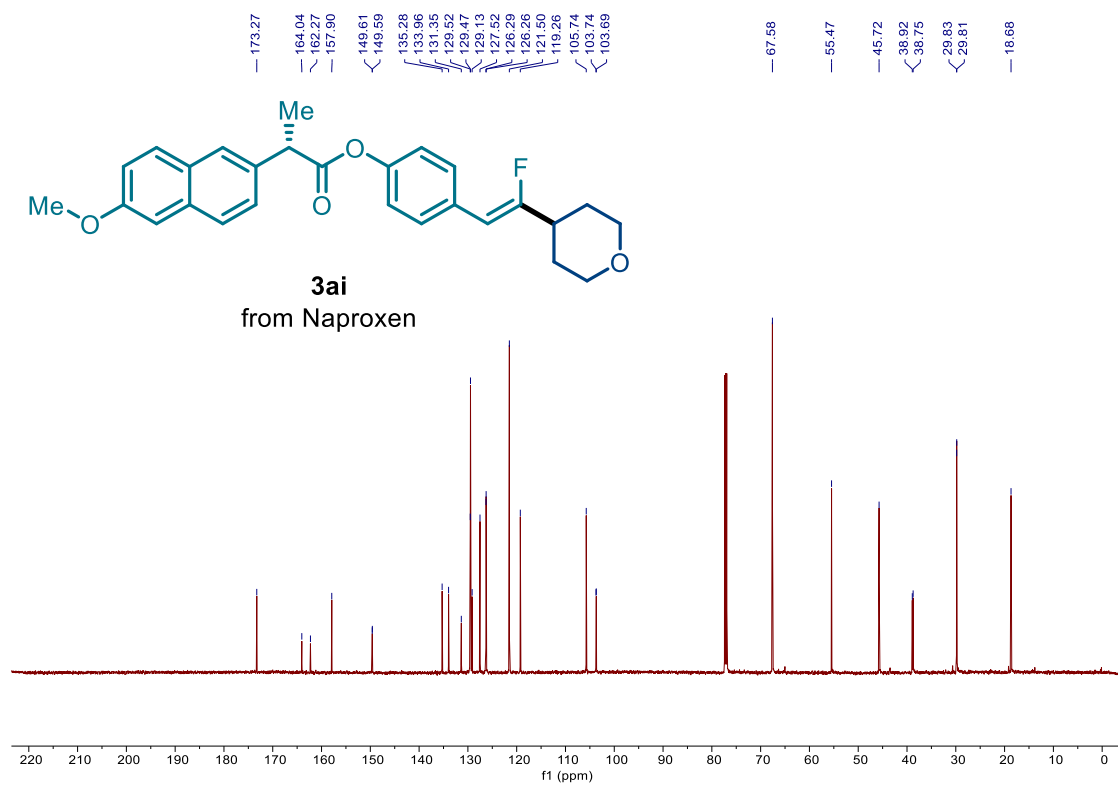
¹⁹F NMR of Compound 3ah (375 MHz, Chloroform-*d*):



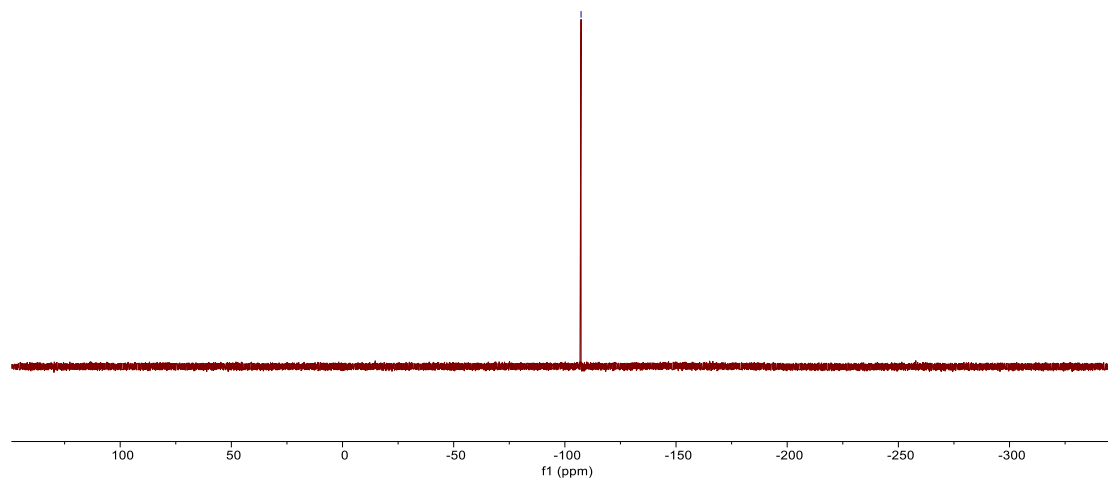
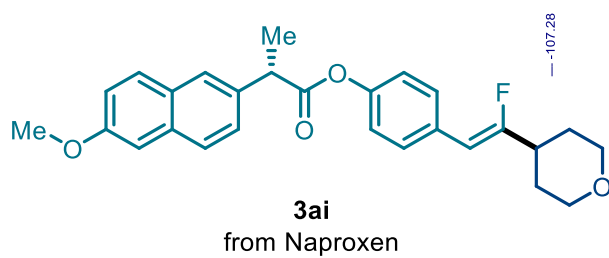
¹H NMR of Compound 3ai (600 MHz, Chloroform-*d*):



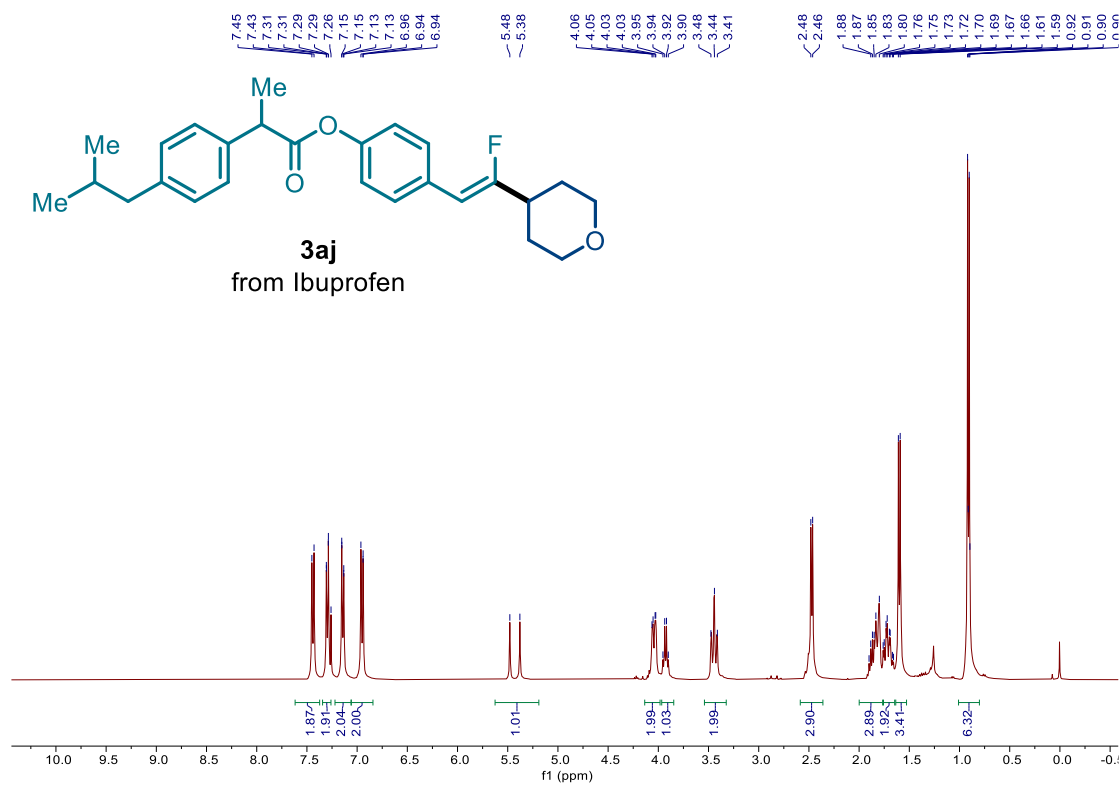
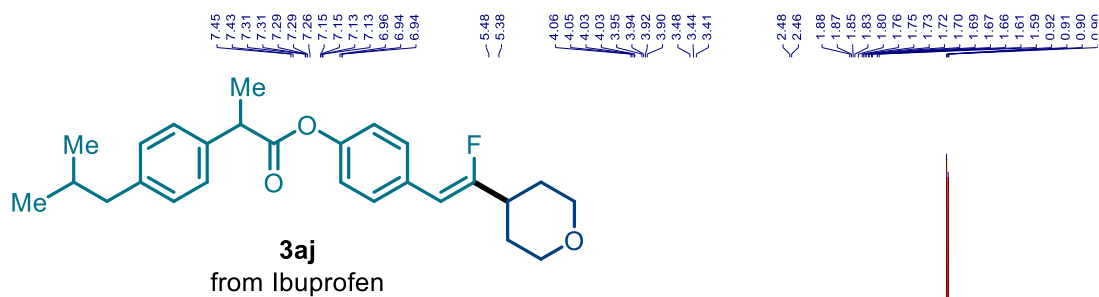
¹³C NMR of Compound 3ai (150 MHz, Chloroform-*d*):



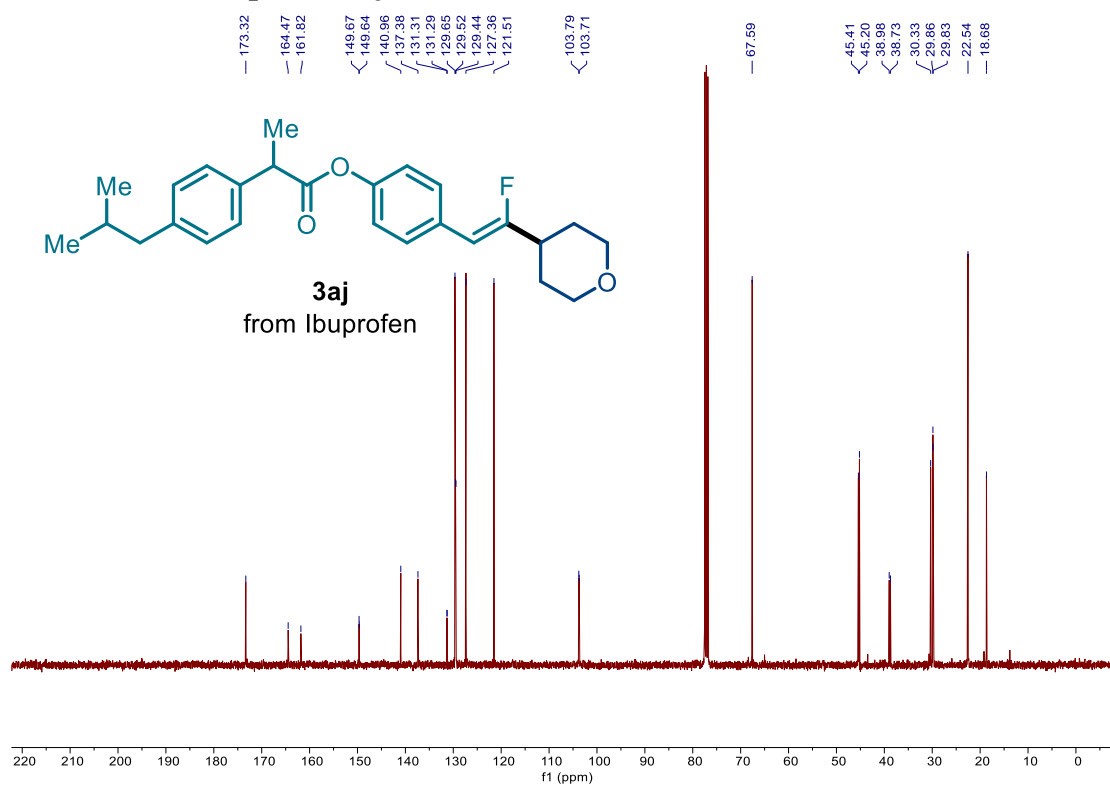
¹⁹F NMR of Compound 3ai (375 MHz, Chloroform-*d*):



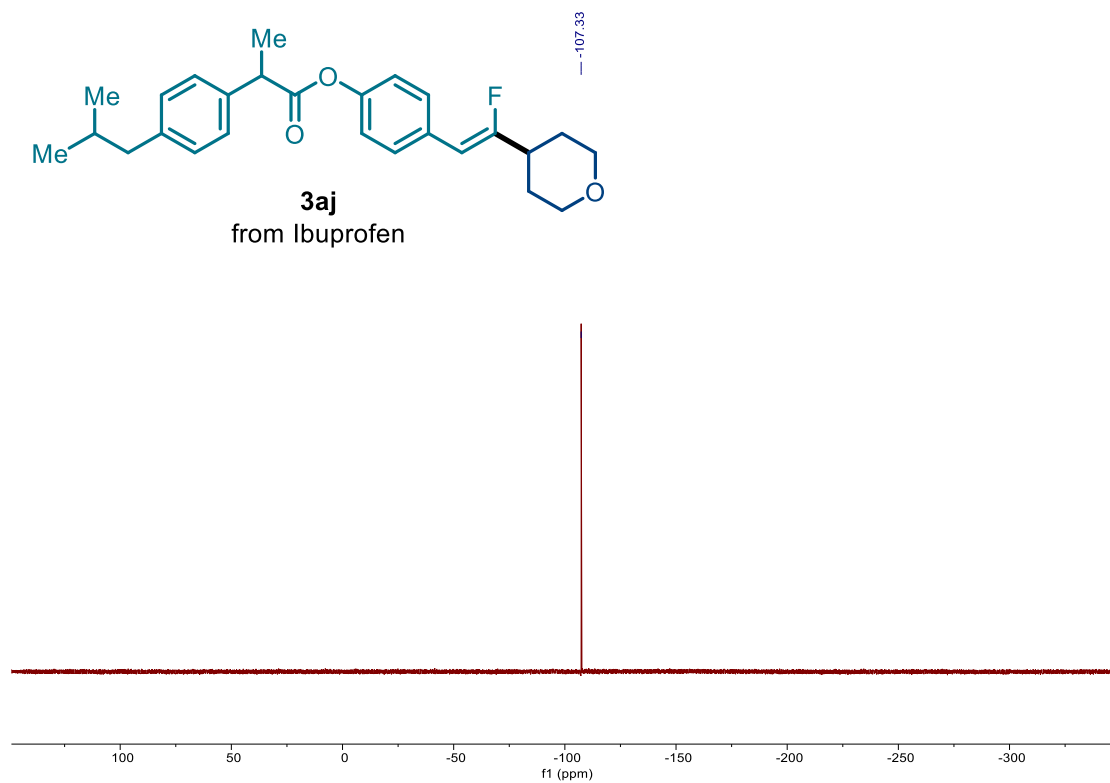
¹H NMR of Compound 3aj (400 MHz, Chloroform-*d*):



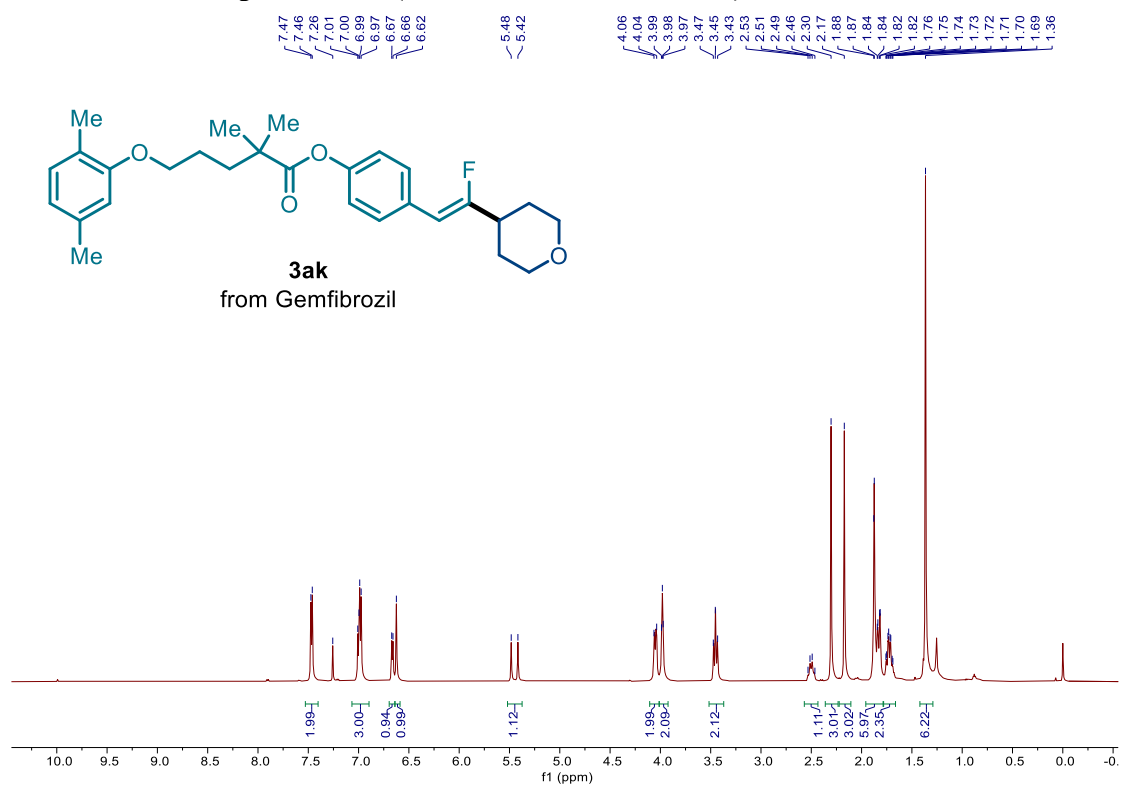
¹³C NMR of Compound 3aj (150 MHz, Chloroform-*d*):



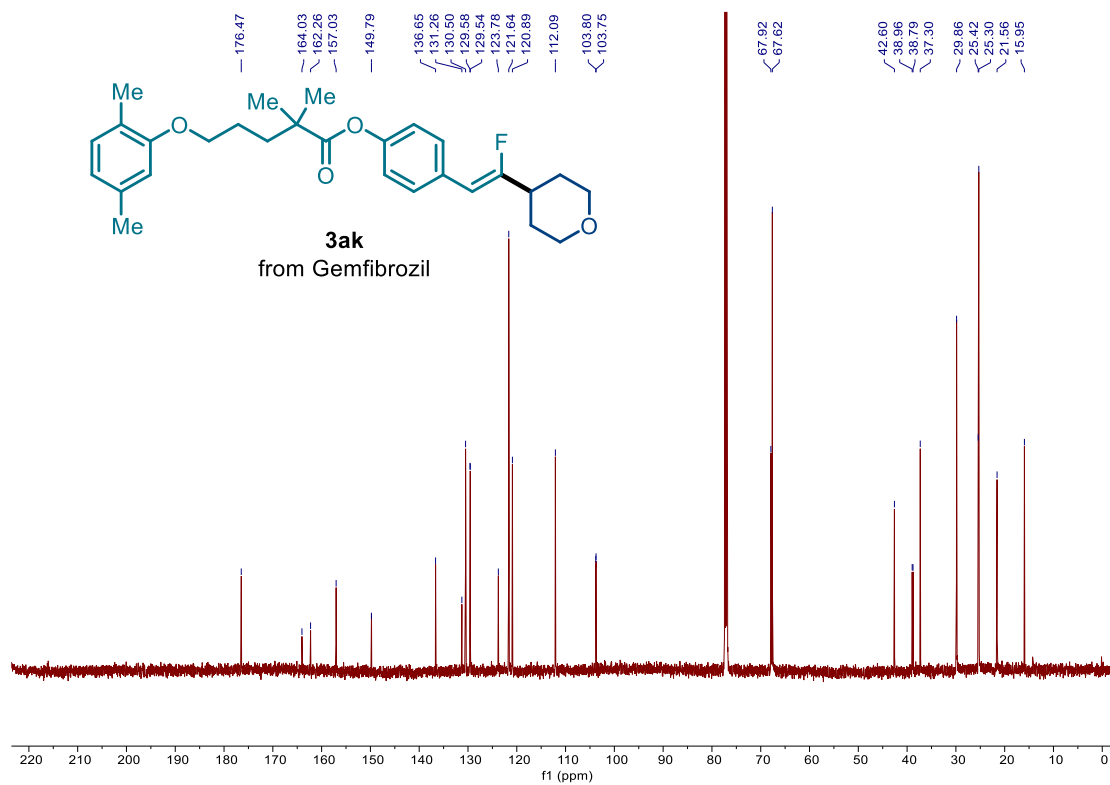
¹⁹F NMR of Compound 3aj (375 MHz, Chloroform-*d*):



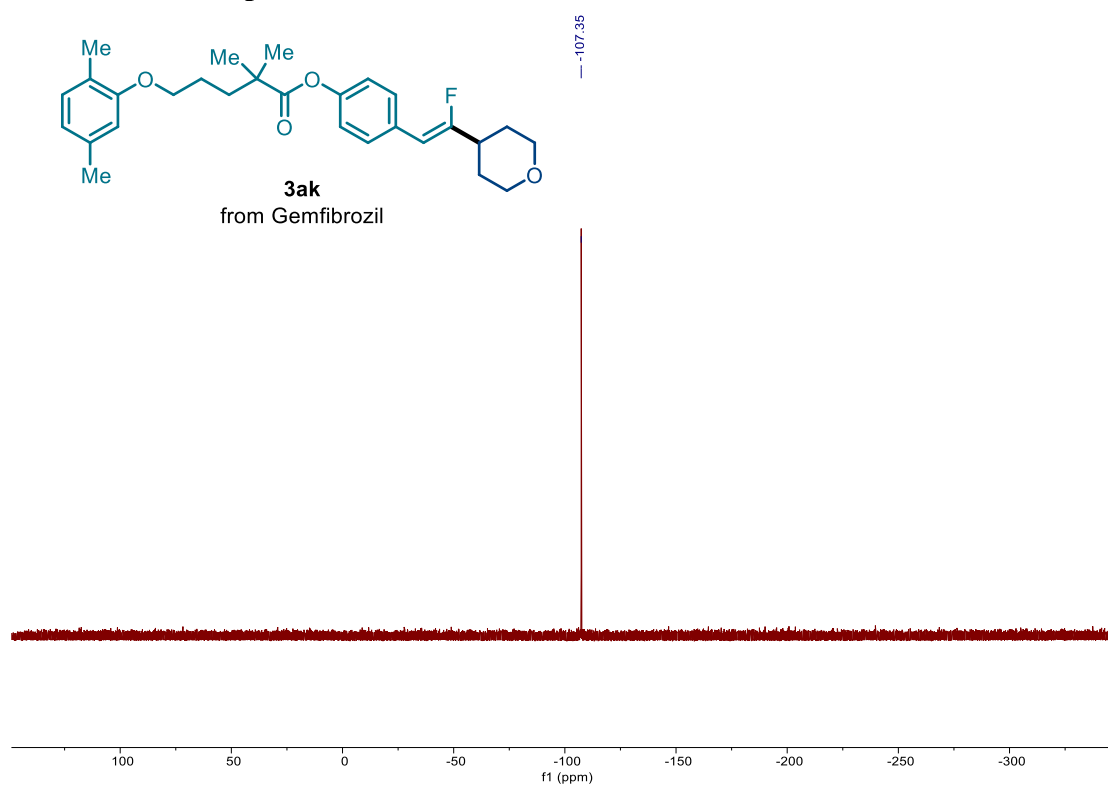
¹H NMR of Compound 3ak (600 MHz, Chloroform-*d*):



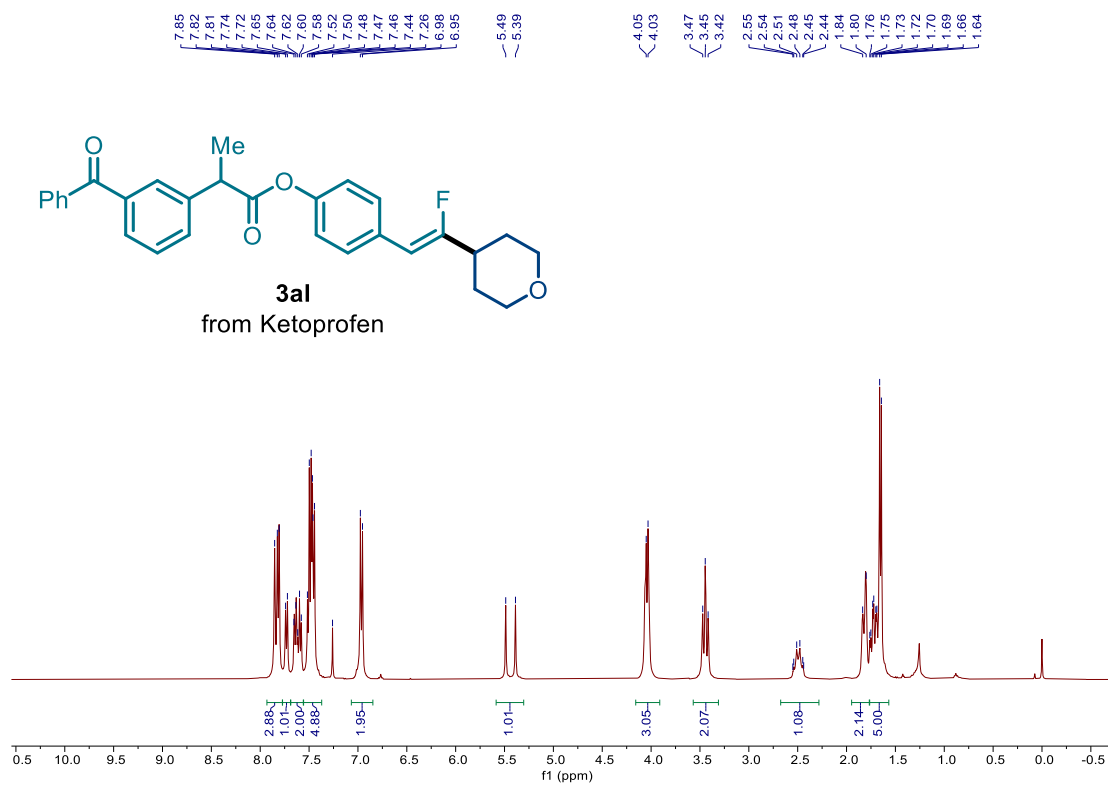
¹³C NMR of Compound 3ak (150 MHz, Chloroform-*d*):



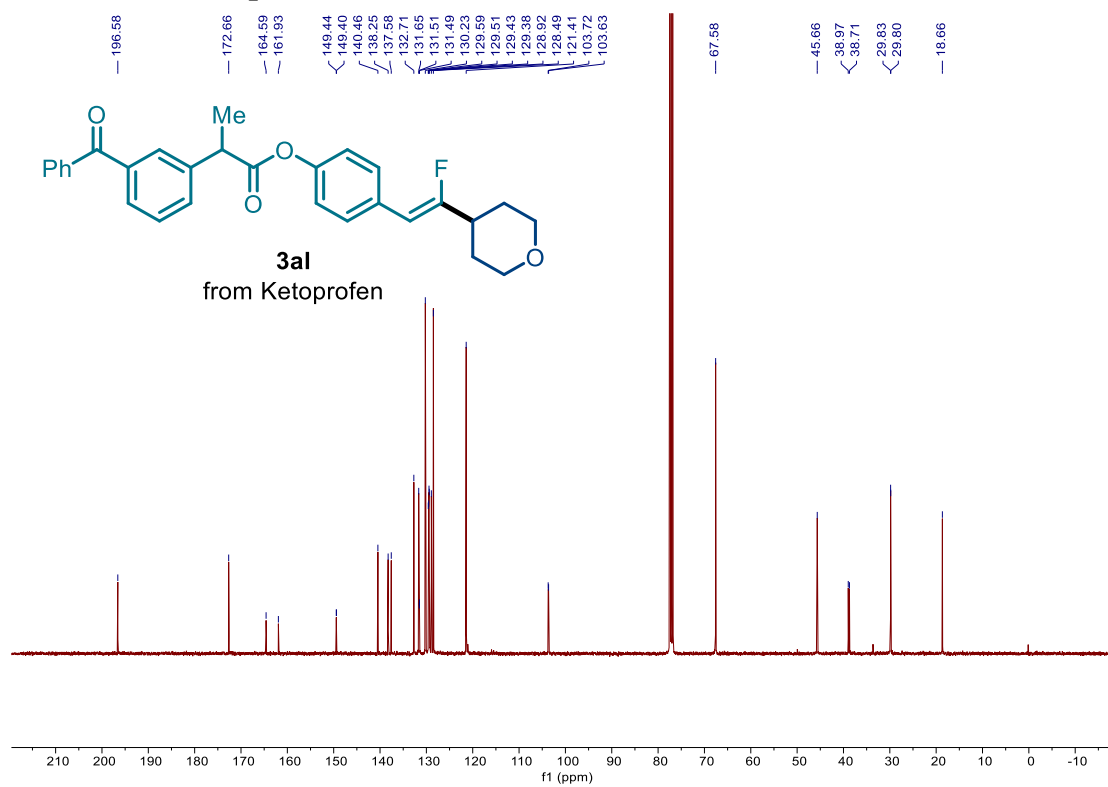
¹⁹F NMR of Compound 3ak (375 MHz, Chloroform-*d*):



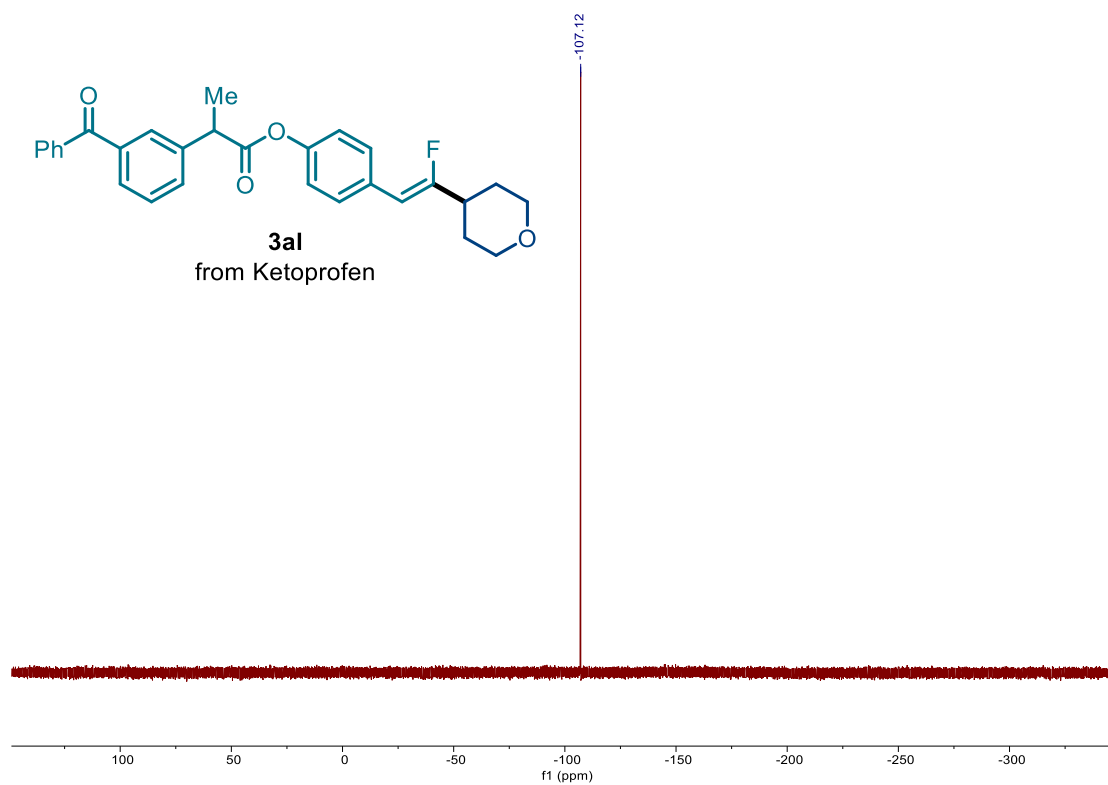
¹H NMR of Compound 3al (400 MHz, Chloroform-*d*):



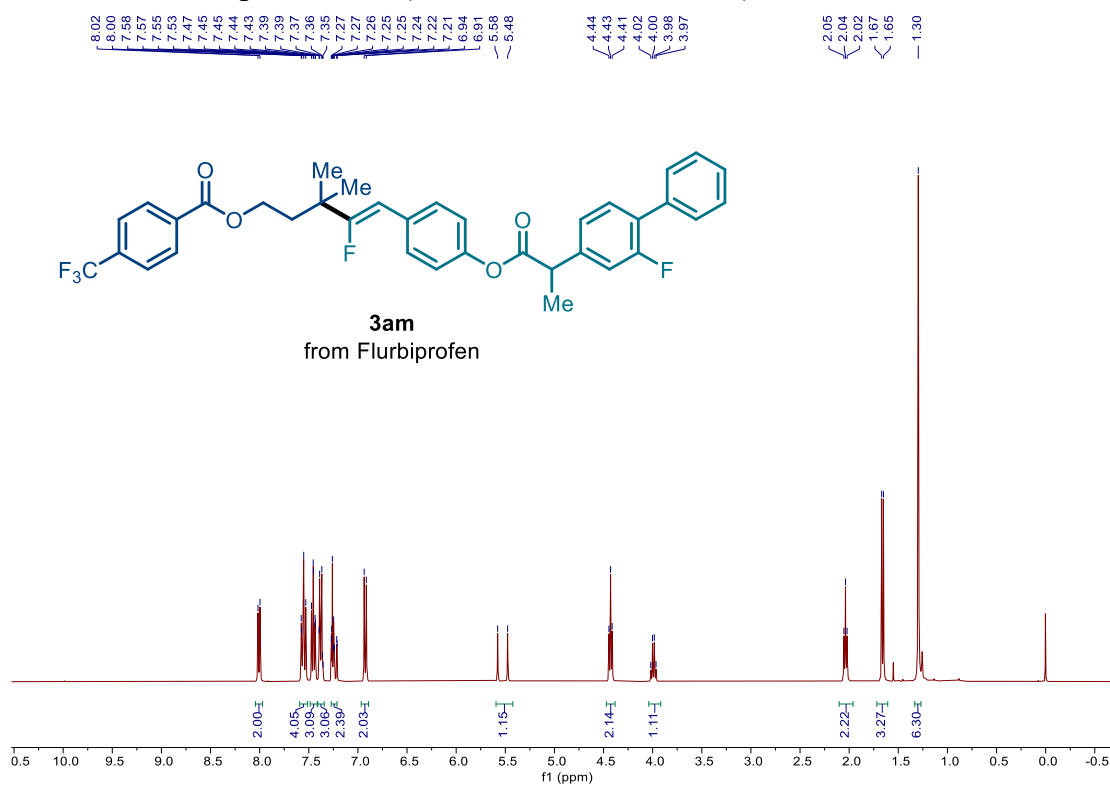
¹³C NMR of Compound 3aI (100 MHz, Chloroform-*d*):



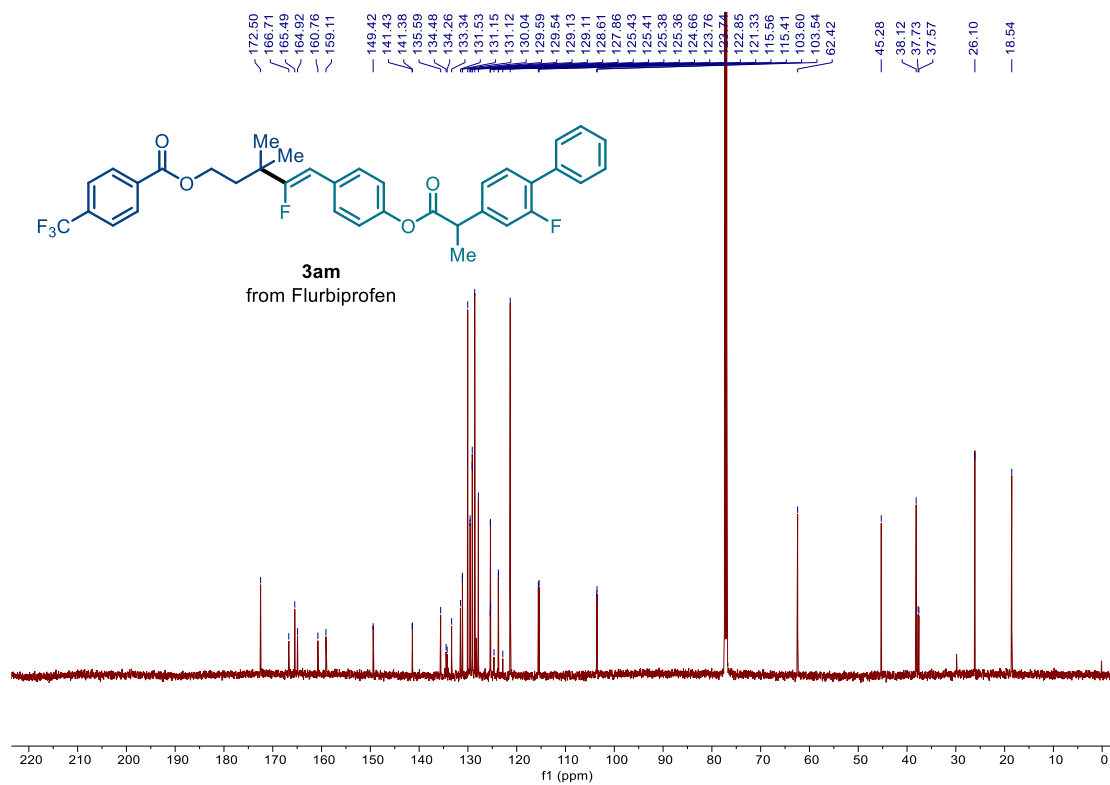
¹⁹F NMR of Compound 3aI (375 MHz, Chloroform-*d*):



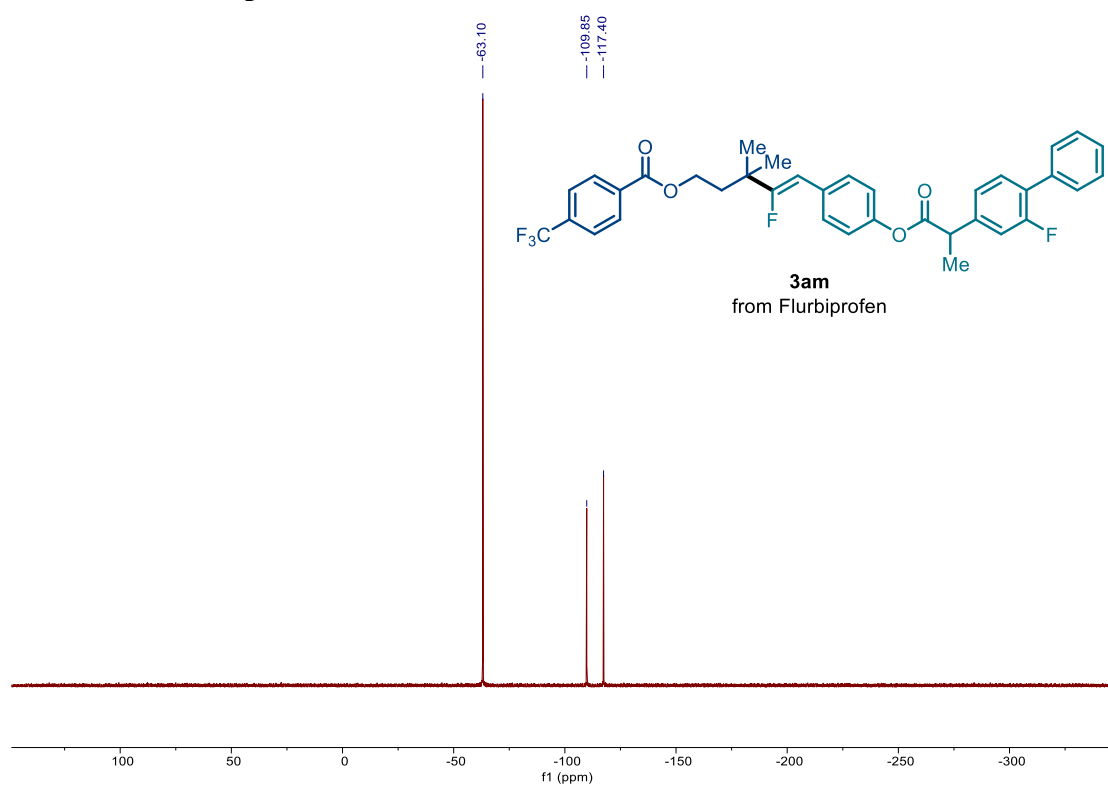
¹H NMR of Compound 3am (400 MHz, Chloroform-*d*):



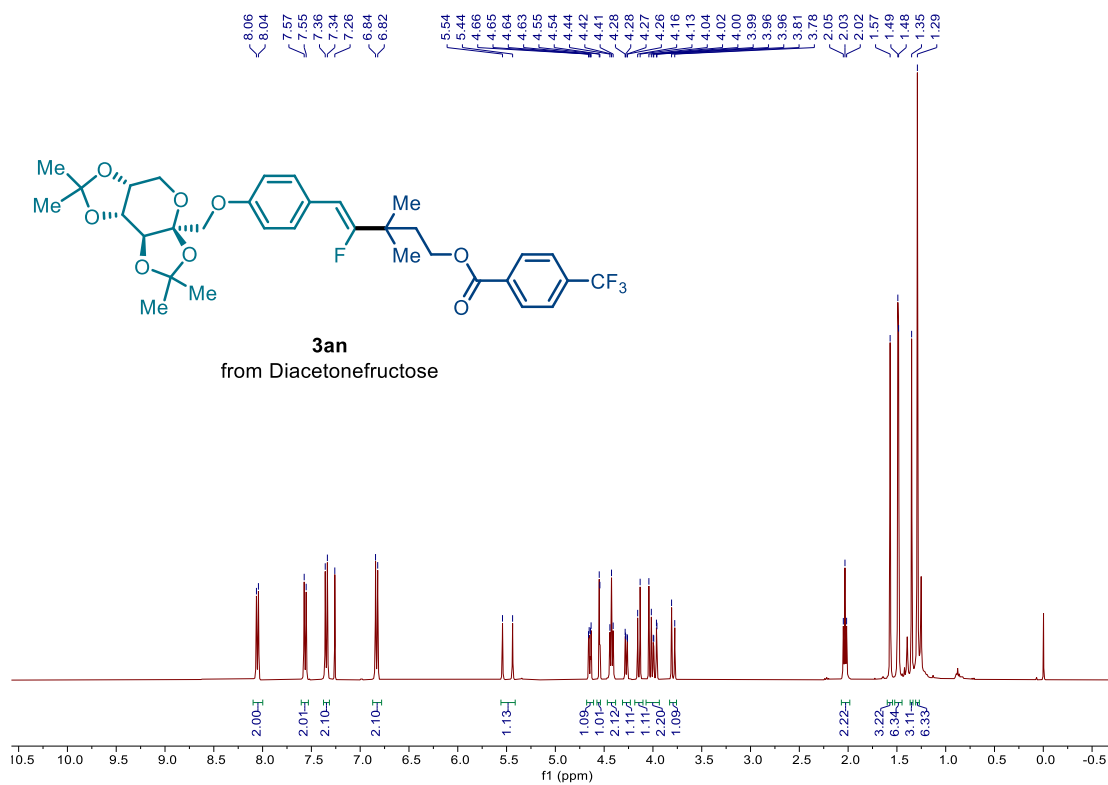
¹³C NMR of Compound 3am (150 MHz, Chloroform-*d*):



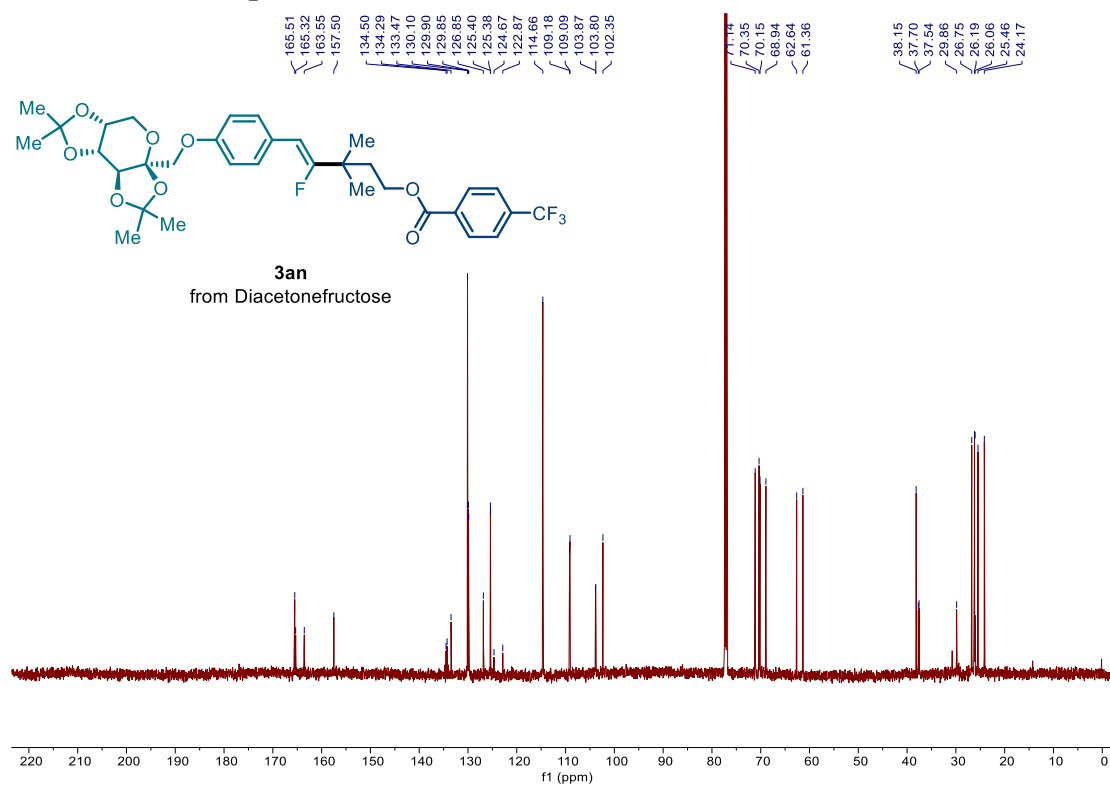
¹⁹F NMR of Compound 3am (375 MHz, Chloroform-*d*):



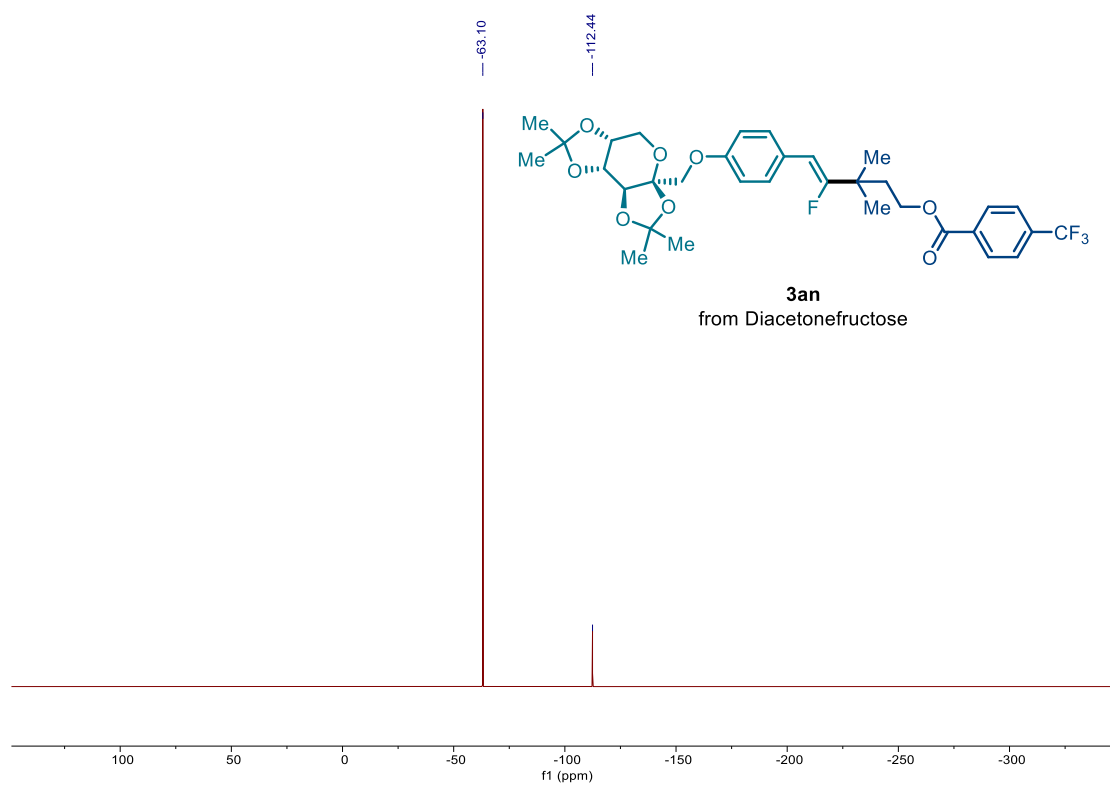
¹H NMR of Compound 3an (400 MHz, Chloroform-*d*):



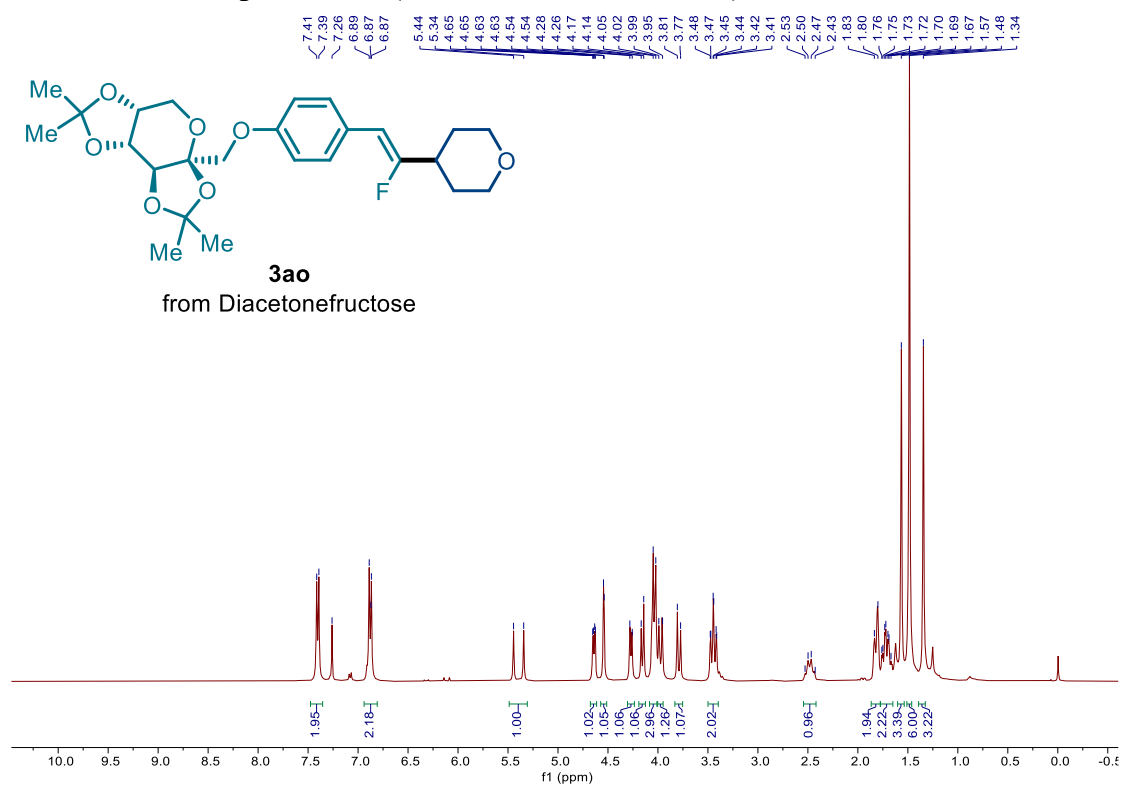
¹³C NMR of Compound 3an (150 MHz, Chloroform-*d*):



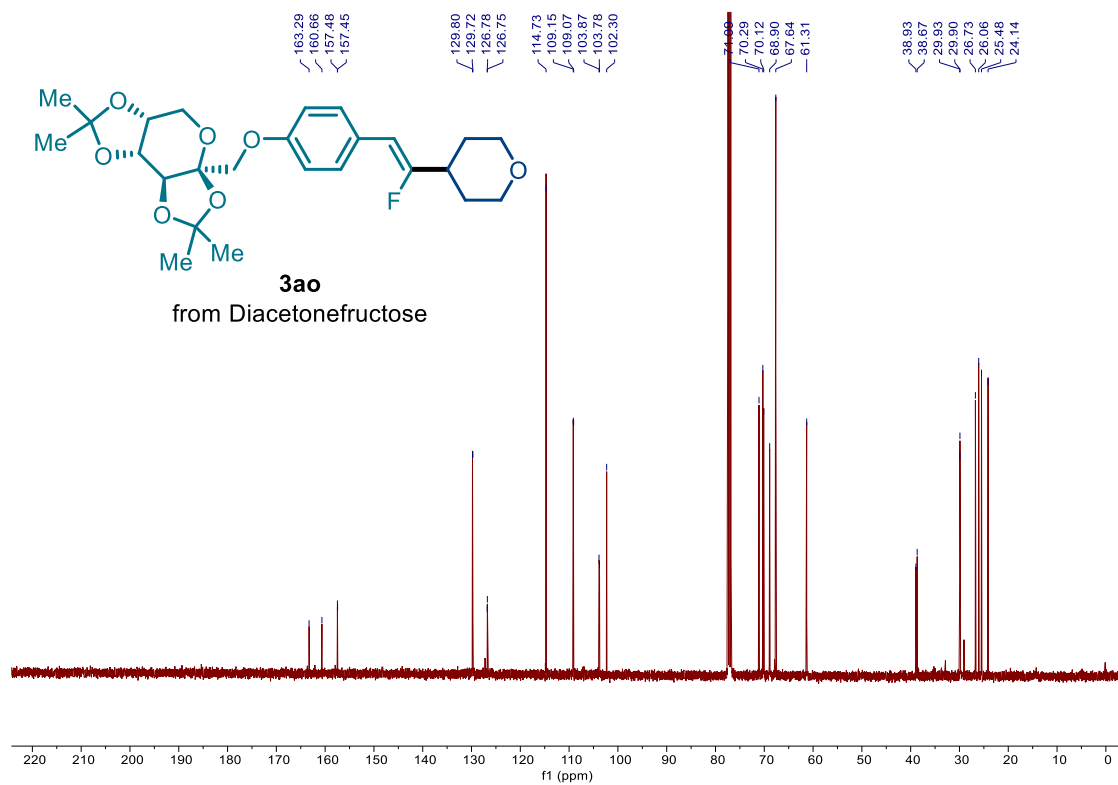
¹⁹F NMR of Compound 3an (375 MHz, Chloroform-*d*):



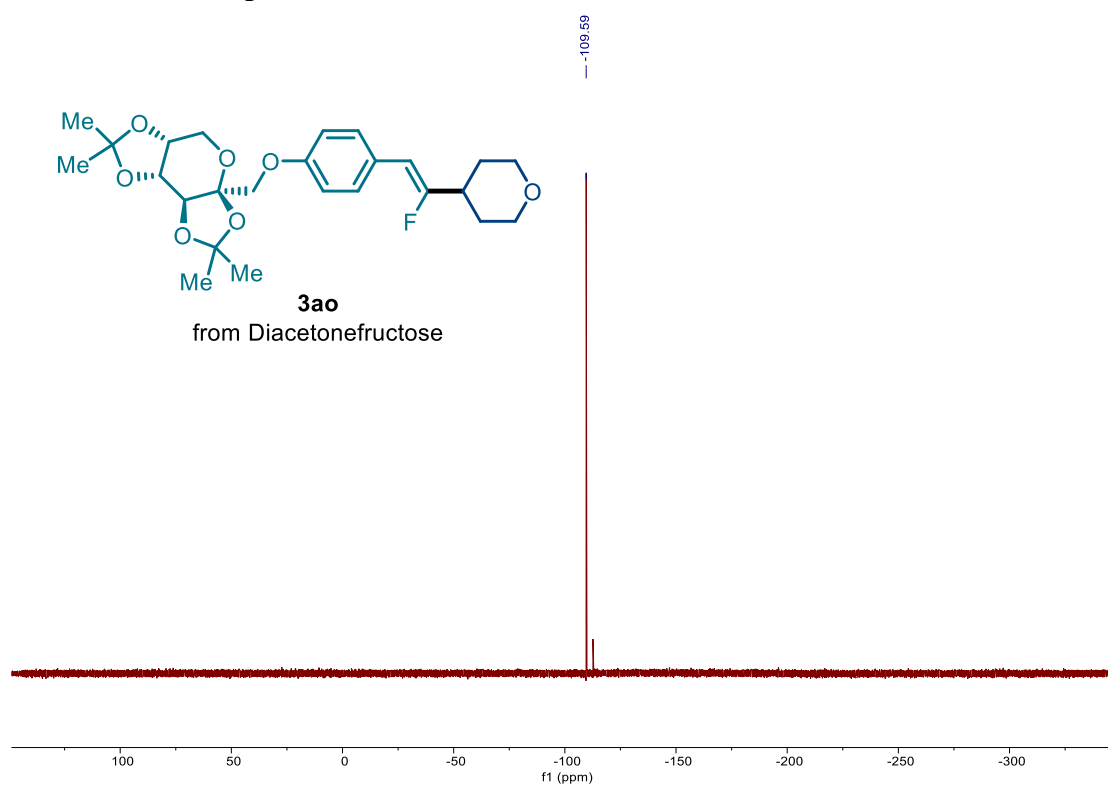
¹H NMR of Compound 3ao (400 MHz, Chloroform-*d*):



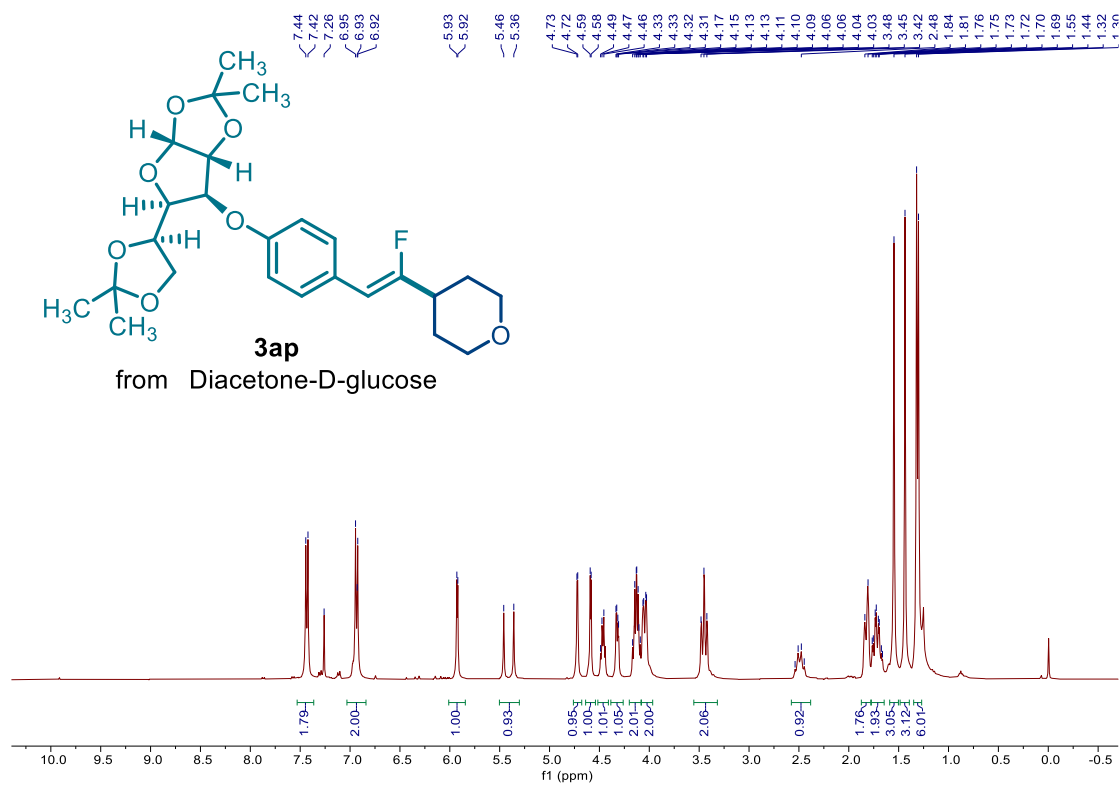
¹³C NMR of Compound 3ao (100 MHz, Chloroform-*d*):



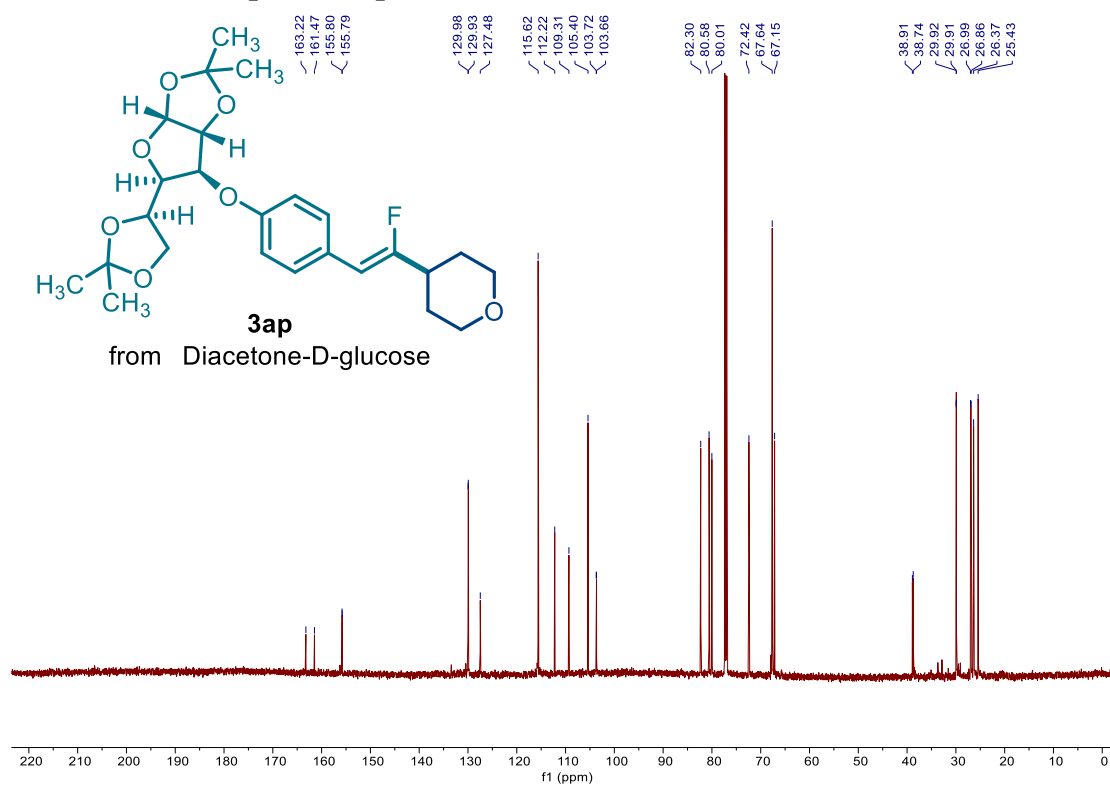
¹⁹F NMR of Compound 3ao (375 MHz, Chloroform-*d*):



¹H NMR of Compound 3ap (400 MHz, Chloroform-*d*):



¹³C NMR of Compound 3ap (150 MHz, Chloroform-*d*):



¹⁹F NMR of Compound 3ap (375 MHz, Chloroform-*d*):

