

Supporting Information

For

Visible light-promoted umpolung coupling of aryl tri-/difluoroethanones with 2-alkenylpyridines

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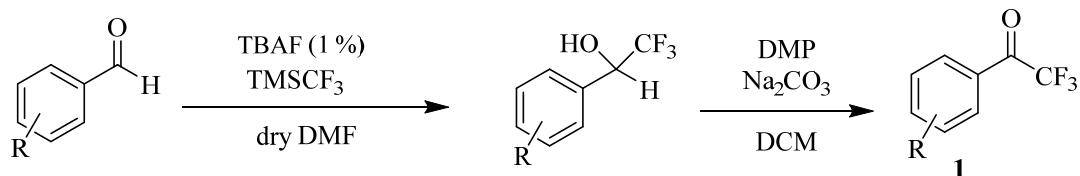
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1. General remarks

¹H NMR spectra were recorded on 400 or 600 MHz (100 or 150 MHz for ¹³C NMR, 564 MHz for ¹⁹F NMR) agilent NMR spectrometer with CDCl₃ as the solvent and tetramethylsilane (TMS) as the internal standard. Chemical shifts were reported in parts per million (ppm, δ scale) downfield from TMS at 0.00 ppm and referenced to the CDCl₃ at 7.26 ppm (for ¹H NMR) or 77.16 ppm (for ¹³C NMR). ¹⁹F NMR chemical shifts were determined relative to CFCl₃ at δ 0.00 ppm. HRMS was recorded on a GCT PremierTM(CI) Mass Spectrometer. Infrared (FT-IR) spectra were recorded on a Varian 1000FT-IR, ν_{max} in cm⁻¹. Melting points were measured using SGW, X-4B and values are uncorrected. All commercially available reagents and solvents were used as received unless otherwise specified.

2. General procedure for synthesis of substrates

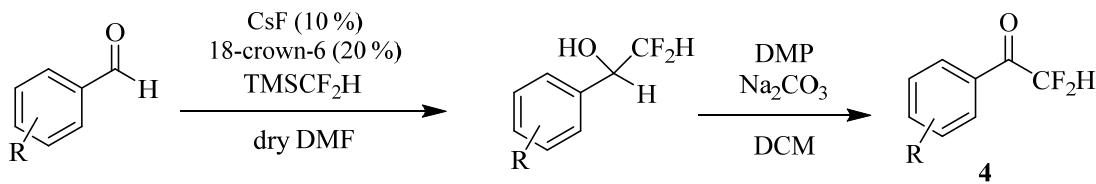
Scheme S1. Synthesis of aryl trifluoroethanones



Step 1: To a solution of aldehyde (5 mmol) in DMF (5 mL) in a 25 mL round-bottom flask equipped, under nitrogen atmosphere, TMSCF₃ (6.5 mmol, 1.3 equiv) was added and the mixture was stirred in an ice bath. After approximately 10 min, TBAF (1 M in THF, 0.05 mmol, 0.05 mL, 0.01 equiv) was added dropwise via a syringe. After 10 min, the ice bath was removed and the solution was stirred for approximately 6 h at room temperature. To cleave the silyl ether intermediate, the reaction mixture was cooled to 0 °C in an ice bath and after 10 min; water and TBAF (1 M in THF, 0.5 mL, 0.5 mmol, 0.1 equiv) were added. The ice bath was removed and the reaction mixture was stirred at room temperature. Finally, the mixture was extracted with ethyl acetate (20 mL \times 3). The organic phase was washed with brine and then dried over anhydrous Na₂SO₄. After filtration and evaporation under vacuum, the residue was subjected to silica gel column chromatography using hexane/ethyl acetate as eluent to give trifluoromethyl alcohols. (*J. Org. Chem.*, **2012**, *77*, 8131–8141)

Step 2: To a solution of the α -CF₃ alcohol (5 mmol, 1 equiv) in DCM (30 mL), was added DMP (17.5 mmol, 3.5 equiv) and NaCO₃ (20 mmol, 4 equiv). The solution was stirred at room temperature for 3 h. Then water was added and the obtained suspension was stirred for an additional hour, the mixture was extracted with DCM (20 mL \times 3). The organic phase was washed with brine and then dried over anhydrous Na₂SO₄. The solvent was removed *in vacuo* by rotary evaporation in a room-temperature water bath to give α -CF₃ ketones **1** (yield up to 97 %). (*J. Org. Chem.*, **1989**, *54*, 661–668; *Tetrahedron Lett.*, **2013**, *54*, 4483–4486)

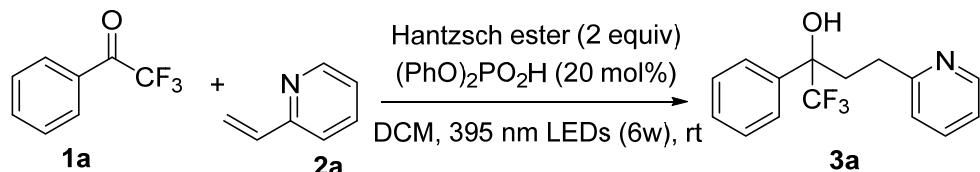
Scheme S2. Synthesis of aryl difluoroethanones



Step 1: Under argon atmosphere, CsF (0.5 mmol, 10 mol %) and 18-crown-6 (1.0 mmol, 20 mol %) were added to a solution of aldehyde (5.0 mmol) in dry DMF (3 mL). TMSCF₂H (6.5 mmol, 1.3 equiv) was added, and the mixture was stirred at room temperature overnight. Subsequently, HCl aq. (1.0 M, 3.5 mL) was added and the solution was stirred for another 1 h. Finally, the mixture was extracted with ethyl acetate (20 mL × 3). The organic phase was washed with brine and then dried over anhydrous Na₂SO₄. After filtration and evaporation under vacuum, the residue was subjected to silica gel column chromatography using hexane/ethyl acetate as eluent to give difluoromethyl alcohols (yield up to 97 %). (*Angew. Chem. Int. Ed.*, **2016**, *55*, 12632–12636)

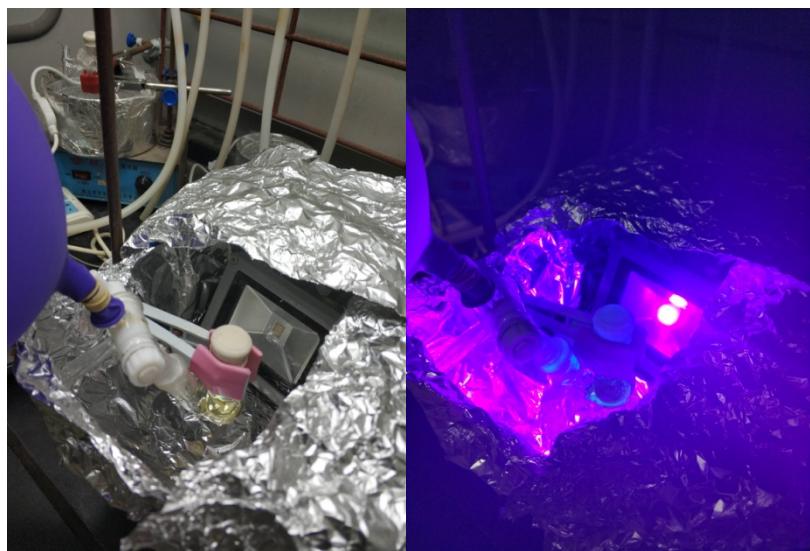
Step 2: Same as scheme 1- step 2.

3. Typical experimental procedure



To a suspension of **1a** (52.2 mg, 0.30 mmol), Hantzsch ester (101mg, 0.40 mmol) and (PhO)₂PO₂H (10 mg, 20 mol%) in DCM (2 mL) was added **2a** (21 mg, 0.20 mmol) at rt. The resulting mixture was stirred upon 6W 395 nm LEDs irradiation under argon balloon. After the reaction was finished, the solvent was removed under reduced pressure and the residue was purified by flash column chromatography on silica gel to give **3a** as a white solid (53.0 mg, 95% yield).

Reaction Setup:



Photographs of the 6w 395 nm LEDs and reaction vessel.

4. UV-vis absorption spectra

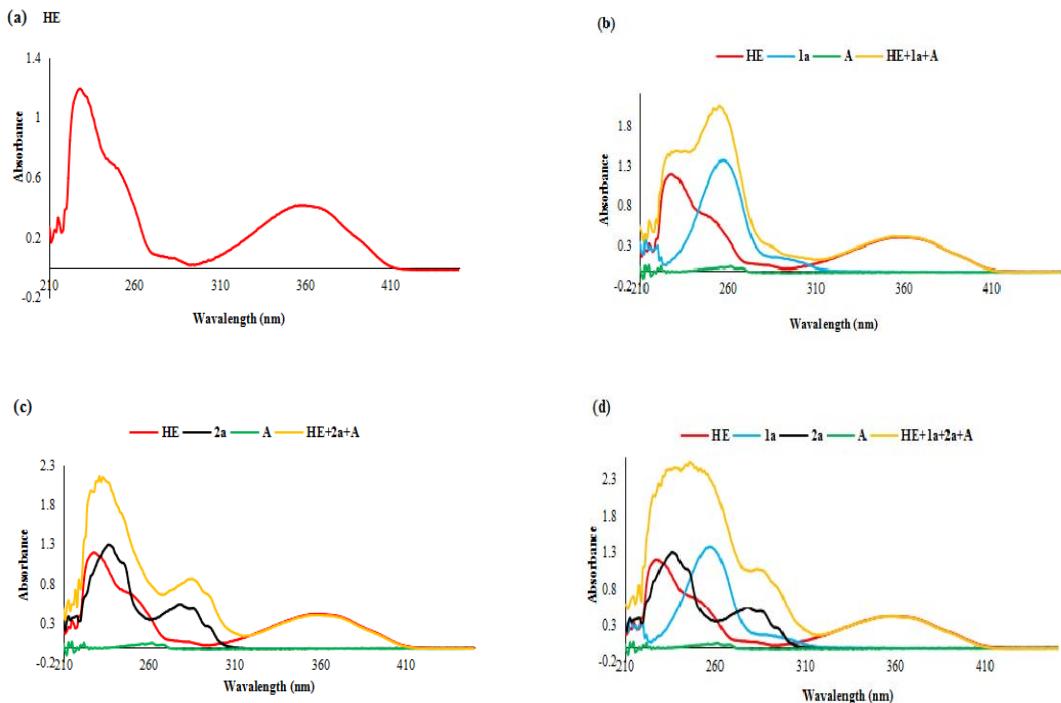


Figure S1. Optical absorption spectra recorded in DCE in 1 cm path quartz cuvettes using a Shimadzu UV-2600 UV-vis spectrophotometer. a) UV-vis absorption spectrum of HE, [HE] = 0.1 μ M. b) UV-vis absorption spectra of HE, α , α , α -trifluoro-1-phenylethan-1-one (**1a**), $(\text{PhO})_2\text{PO}_2\text{H}$ (**A**), and the mixture of HE, **1a** and **A**, [HE] = 0.1 μ M, [**1a**] = 0.1 μ M, [**A**] = 0.02 μ M. c) UV-vis absorption spectra of HE, 2-vinylpyridine (**2a**), $(\text{PhO})_2\text{PO}_2\text{H}$ (**A**), and the mixture of HE, **2a** and **A**, [HE] = 0.1 μ M, [**2a**] = 0.1 μ M, [**A**] = 0.02 μ M. d) UV-vis absorption spectra of HE, α , α , α -trifluoro-1-phenylethan-1-one (**1a**), 2-vinylpyridine (**2a**), $(\text{PhO})_2\text{PO}_2\text{H}$ (**A**), and the mixture of HE, **1a**, **2a** and **A**, [HE] = 0.1 μ M, [**1a**] = 0.1 μ M, [**2a**] = 0.1 μ M, [**A**] = 0.02 μ M.

α -trifluoro-1-phenylethan-1-one (**1a**), 2-vinylpyridine (**2a**), $(\text{PhO})_2\text{PO}_2\text{H}$ (**A**), and the mixture of HE, **1a**, **2a** and **A**, $[\text{HE}] = 0.1 \mu\text{M}$, $[\text{1a}] = 0.1 \mu\text{M}$, $[\text{2a}] = 0.1 \mu\text{M}$, $[\text{A}] = 0.02 \mu\text{M}$.

5. Luminescence quenching experiments

Emission intensities were recorded using LS55 Luminescence Spectrometer for all experiments. All HE solutions were excited at 390 nm and the emission intensity was collected at 440-460nm. In a typical experiment, the DCE solution of HE ($0.1 \mu\text{M}$) and $(\text{PhO})_2\text{POOH}$ ($0.02 \mu\text{M}$) was added the appropriate amount of α , α , α -trifluoro-1-phenylethan-1-one (**1a**) or 2-vinylpyridine (**2a**) in a screw-top 1.0 cm quartz cuvette. After degassing with nitrogen for 10 min, the emission spectra of the samples were collected. The results showed that α , α , α -trifluoro-1-phenylethan-1-one (**1a**) quenched the photoexcited HE* effectively but 2-vinylpyridine (**2a**) did not.

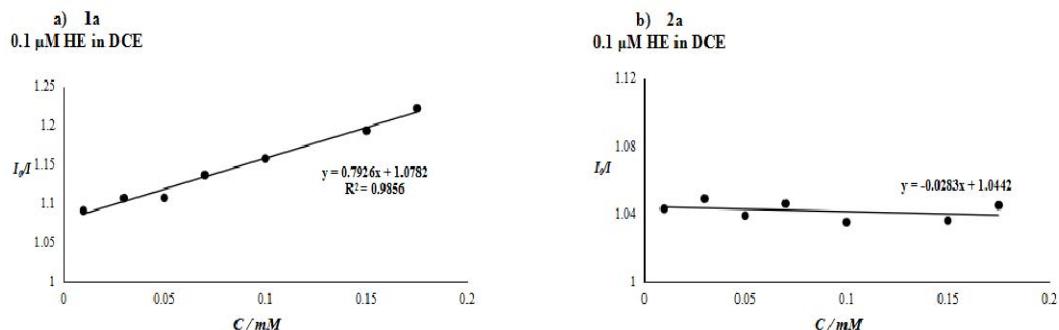
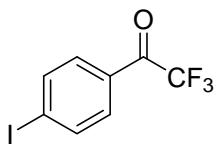


Figure S2. a) HE emission quenching by α , α , α -trifluoro-1-phenylethan-1-one (**1a**). b) HE emission quenching by 2-vinylpyridine (**2a**).

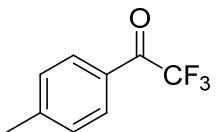
6. References for known substrates

Entry	References	Compounds
1	W. Wu, et al. <i>Chem. Eur. J.</i> , 2016 , 22, 16455-16458.	1e, 1f, 1g
2	D. J. Leng, et al. <i>Org. Biomol. Chem.</i> , 2016 , 14, 1531-1535.	4a, 4b, 4c, 4d, 4f
3	E. Schmitt, et al. <i>Org. Lett.</i> , 2015 , 17, 4510-4513.	4e

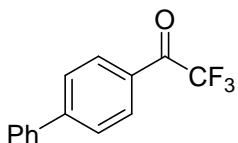
7. Characterization of the substrates and products



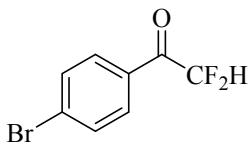
2,2,2-Trifluoro-1-(4-iodophenyl)ethan-1-one (1e): ^1H NMR (400 MHz, CDCl_3) δ 7.93 (d, $J = 8.3$ Hz, 2H), 7.76 (d, $J = 8.1$ Hz, 2H); ^{19}F NMR (564 MHz, CDCl_3) δ -71.61 (s, 3F).



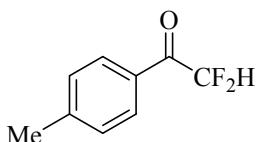
2,2,2-Trifluoro-1-(p-tolyl)ethan-1-one (1f): ^1H NMR (400 MHz, CDCl_3) δ 7.97 (d, $J = 7.8$ Hz, 2H), 7.34 (d, $J = 8.0$ Hz, 2H), 2.46 (s, 3H); ^{19}F NMR (564 MHz, CDCl_3) δ -71.38 (s, 3F).



1-([1,1'-Biphenyl]-4-yl)-2,2,2-trifluoroethan-1-one (1g): ^1H NMR (400 MHz, CDCl_3) δ 8.16 (d, $J = 8.0$ Hz, 2H), 7.77 (d, $J = 8.2$ Hz, 2H), 7.65 (d, $J = 7.4$ Hz, 2H), 7.51 (t, $J = 7.3$ Hz, 2H), 7.48 – 7.42 (t, 1H); ^{19}F NMR (564 MHz, CDCl_3) δ -71.36 (s, 3F).

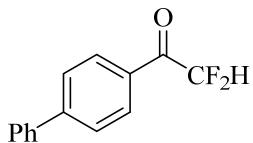


1-(4-Bromophenyl)-2,2-difluoroethan-1-one (4a): ^1H NMR (400 MHz, CDCl_3) δ 7.91 (d, $J = 8.2$ Hz, 2H), 7.65 (d, $J = 8.4$ Hz, 2H), 6.24 (t, $J = 53.4$ Hz, 1H); ^{19}F NMR (564 MHz, CDCl_3) δ -122.02 (d, $J_{H-F} = 53.3$, 2F).

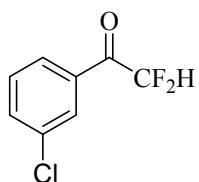


2,2-Difluoro-1-(p-tolyl)ethan-1-one (4b): ^1H NMR (400 MHz, CDCl_3) δ 7.96 (d, $J =$

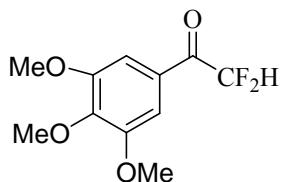
7.9 Hz, 2H), 7.31 (d, J = 7.9 Hz, 2H), 6.28 (t, J = 53.6 Hz, 1H), 2.43 (s, 3H); ^{19}F NMR (564 MHz, CDCl_3) δ -122.10 (d, $J_{\text{H}-\text{F}}$ = 53.6 Hz, 2F).



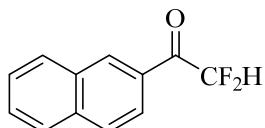
1-([1,1'-Biphenyl]-4-yl)-2,2-difluoroethan-1-one (4c): ^1H NMR (400 MHz, CDCl_3) δ 8.16 (d, J = 8.1 Hz, 2H), 7.75 (d, J = 8.2 Hz, 2H), 7.65 (d, J = 7.4 Hz, 2H), 7.50 (t, J = 7.3 Hz, 2H), 7.47 – 7.41 (m, 1H), 6.33 (t, J = 53.5 Hz, 1H); ^{19}F NMR (564 MHz, CDCl_3) δ -121.77 (d, $J_{\text{H}-\text{F}}$ = 53.6 Hz, 2F).



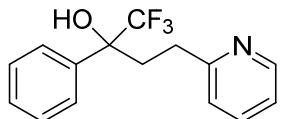
1-(3-Chlorophenyl)-2,2-difluoroethan-1-one (4d): ^1H NMR (600 MHz, CDCl_3) δ 8.03 (s, 1H), 7.96 (d, J = 7.8 Hz, 1H), 7.64 (m, 1H), 7.48 (t, J = 7.9 Hz, 1H), 6.25 (t, J = 53.4 Hz, 1H); ^{19}F NMR (564 MHz, CDCl_3) δ -121.85 (d, $J_{\text{H}-\text{F}}$ = 53.3 Hz, 2F).



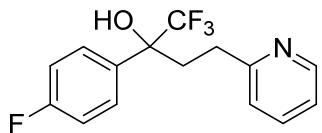
2,2-Difluoro-1-(3,4,5-trimethoxyphenyl) ethan-1-one (4e): ^1H NMR (400 MHz, CDCl_3) δ 7.32 (s, 2H), 6.26 (t, J = 53.6 Hz, 1H), 3.95 (s, 3H), 3.92 (s, 6H); ^{19}F NMR (564 MHz, CDCl_3) δ -120.71 (d, $J_{\text{H}-\text{F}}$ = 53.6 Hz, 2F).



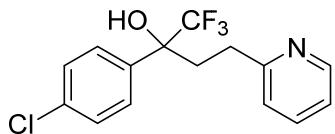
2,2-Difluoro-1-(naphthalen-2-yl) ethan-1-one (4f): ^1H NMR (400 MHz, CDCl_3) δ 8.62 (s, 1H), 8.04 (t, J = 10 – 8 Hz, 1H), 7.97 (d, J = 8.0 Hz, 1H), 7.89 (dd, J = 15.4, 8.4 Hz, 2H), 7.64 (dd, J = 14.9, 7.2 Hz, 1H), 7.58 (t, J = 7.4 Hz, 1H), 6.43 (t, J = 53.5 Hz, 1H); ^{19}F NMR (564 MHz, CDCl_3) δ -121.43 (d, $J_{\text{H}-\text{F}}$ = 53.5 Hz, 2F).



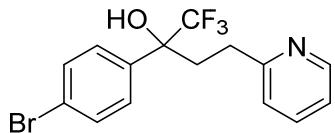
1,1,1-Trifluoro-2-phenyl-4-(pyridin-2-yl)butan-2-ol (3a): White solid; m.p. 115–117 °C; 95% yield (53 mg); ¹H NMR (600 MHz, CDCl₃) δ 9.38 (s, 1H), 8.48 (d, *J* = 4.7 Hz, 1H), 7.73 (d, *J* = 7.8 Hz, 2H), 7.60 (t, *J* = 7.6 Hz, 1H), 7.40 (t, *J* = 7.6 Hz, 2H), 7.34 (t, *J* = 7.2 Hz, 1H), 7.19 – 7.14 (m, 1H), 7.12 (d, *J* = 7.8 Hz, 1H), 2.94 (dd, *J* = 16.8, 8.8 Hz, 1H), 2.78 – 2.67 (m, 2H), 2.59 (dd, *J* = 14.9, 9.8 Hz, 1H); ¹³C NMR (150 MHz, CDCl₃) δ 160.1, 147.7, 139.0, 137.6, 128.20, 128.1, 127.3, 124.4 (q, *J*_{C-F} = 285.9 Hz), 123.6, 121.60, 76.5 (q, *J*_{C-F} = 27.4 Hz), 32.2, 31.1; ¹⁹F NMR (564 MHz, CDCl₃) δ -79.68 (s, 3F); FT-IR (thin film, KBr): ν (cm⁻¹) 3061, 1481, 1450, 702; HRMS (CI) calcd C₁₅H₁₅NOF₃ [M + H]⁺: 282.1106, found: 282.1113.



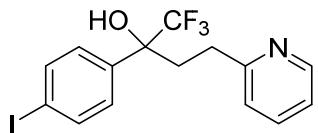
1,1,1-Trifluoro-2-(4-fluorophenyl)-4-(pyridin-2-yl)butan-2-ol (3b): White solid; m.p. 107–109 °C; 54% yield (32 mg); ¹H NMR (600 MHz, CDCl₃) δ 9.49 (s, 1H), 8.49 (d, *J* = 4.9 Hz, 1H), 7.68 (dd, *J* = 8.4, 5.6 Hz, 2H), 7.65 – 7.60 (m, 1H), 7.21 – 7.17 (m, 1H), 7.14 (d, *J* = 7.8 Hz, 1H), 7.07 (t, *J* = 8.7 Hz, 2H), 2.97 (dd, *J* = 16.9, 8.8 Hz, 1H), 2.79 – 2.63 (m, 2H), 2.58 (dd, *J* = 14.9, 9.7 Hz, 1H); ¹³C NMR (150 MHz, CDCl₃) δ 162.7 (d, *J*_{C-F} = 246.7 Hz), 160.1, 147.8, 137.7, 134.9 (d, *J*_{C-F} = 2.9 Hz), 129.3 (d, *J*_{C-F} = 8.1 Hz), 126.2 (q, *J*_{C-F} = 285.7 Hz), 123.6, 121.8, 115.1 (d, *J*_{C-F} = 21.3 Hz), 76.3 (q, *J*_{C-F} = 27.7 Hz), 32.1, 31.1; ¹⁹F NMR (376 MHz, CDCl₃) δ -79.95 (s, 3F), -114.75 – -114.86 (m, 1F); FT-IR (thin film, KBr): ν (cm⁻¹) 3062, 2920, 1600, 1509, 828; HRMS (CI) calcd C₁₅H₁₄NOF₄ [M + H]⁺: 300.1012, found: 300.1020.



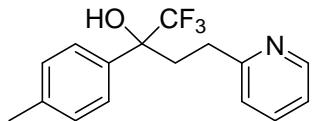
2-(4-Chlorophenyl)-1,1,1-trifluoro-4-(pyridin-2-yl)butan-2-ol (3c): White solid; m.p. 105–106 °C; 94% yield (59 mg); ¹H NMR (400 MHz, CDCl₃) δ 8.49 (d, *J* = 3.9 Hz, 1H), 7.74 – 7.54 (m, 3H), 7.36 (d, *J* = 8.3 Hz, 2H), 7.24 – 7.00 (m, 2H), 2.95 (dd, *J* = 16.5, 7.7 Hz, 1H), 2.82 – 2.47 (m, 3H); ¹³C NMR (150 MHz, CDCl₃) δ 159.8, 147.3, 138.2, 137.6, 134.3, 128.9, 128.5, 126.1 (q, *J*_{C-F} = 285.9 Hz), 123.9, 122.0, 76.3 (q, *J*_{C-F} = 27.7 Hz), 32.3, 30.9; ¹⁹F NMR (564 MHz, CDCl₃) δ -79.85 (s, 3F); FT-IR (thin film, KBr): ν (cm⁻¹) 3097, 2924, 1597, 1458, 821; HRMS (CI) calcd C₁₅H₁₄NOF₃³⁵Cl [M + H]⁺: 316.0716, found: 316.0727.



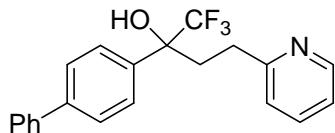
2-(4-Bromophenyl)-1,1,1-trifluoro-4-(pyridin-2-yl)butan-2-ol (3d): White solid; m.p. 137–139 °C; 70% yield (50 mg); ¹H NMR (400 MHz, CDCl₃) δ 9.59 (s, 1H), 8.48 (d, *J* = 4.4 Hz, 1H), 7.63 (t, *J* = 7.7 Hz, 1H), 7.58 (d, *J* = 8.4 Hz, 2H), 7.51 (d, *J* = 8.4 Hz, 2H), 7.23 – 7.16 (m, 1H), 7.14 (d, *J* = 7.8 Hz, 1H), 2.96 (dd, *J* = 16.8, 8.4 Hz, 1H), 2.79 – 2.50 (m, 3H); ¹³C NMR (150 MHz, CDCl₃) δ 159.9, 147.7, 138.3, 137.7, 131.4, 129.3, 126.1 (q, *J*_{C-F} = 285.9 Hz), 123.7, 122.5, 121.7, 76.4 (q, *J*_{C-F} = 27.6 Hz), 32.1, 31.1; ¹⁹F NMR (564 MHz, CDCl₃) δ -79.82 (s, 3F); FT-IR (thin film, KBr): ν (cm⁻¹) 3081, 1489, 1437, 838, 725; HRMS (CI) calcd C₁₅H₁₄NOF₃⁷⁹Br [M + H]⁺: 360.0211, found: 360.0221.



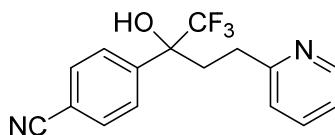
1,1,1-Trifluoro-2-(4-iodophenyl)-4-(pyridin-2-yl)butan-2-ol (3e): White solid; m.p. 150–152 °C; 99% yield (81 mg); ¹H NMR (600 MHz, CDCl₃) δ 9.52 (s, 1H), 8.47 (d, *J* = 4.7 Hz, 1H), 7.72 (d, *J* = 8.5 Hz, 2H), 7.66 – 7.56 (m, 1H), 7.45 (d, *J* = 8.3 Hz, 2H), 7.22 – 7.15 (m, 1H), 7.13 (d, *J* = 7.8 Hz, 1H), 2.95 (dd, *J* = 16.8, 8.4 Hz, 1H), 2.71 (dd, *J* = 16.9, 9.5 Hz, 1H), 2.67 – 2.60 (m, 1H), 2.61 – 2.51 (m, 1H); ¹³C NMR (150 MHz, CDCl₃) δ 159.9, 147.7, 139.0, 137.7, 137.4, 129.5, 126.0 (q, *J*_{C-F} = 285.9 Hz), 123.7, 121.8, 94.4, 76.4 (q, *J*_{C-F} = 27.6 Hz), 32.0, 31.1; ¹⁹F NMR (564 MHz, CDCl₃) δ -79.76 (s, 3F); FT-IR (thin film, KBr): ν (cm⁻¹) 3090, 2922, 1596, 1457, 723; HRMS (CI) calcd C₁₅H₁₄NOF₃I [M + H]⁺: 408.0072, found: 408.0085.



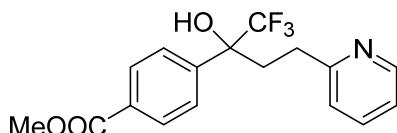
1,1,1-Trifluoro-4-(pyridin-2-yl)-2-(p-tolyl)butan-2-ol (3f): White solid; m.p. 120–122 °C; 83% yield (49 mg); ¹H NMR (400 MHz, CDCl₃) δ 9.27 (s, 1H), 8.49 (d, *J* = 4.4 Hz, 1H), 7.68 – 7.50 (m, 3H), 7.23 – 7.08 (m, 4H), 2.94 (dd, *J* = 16.7, 8.6 Hz, 1H), 2.84 – 2.63 (m, 2H), 2.56 (dd, *J* = 13.9, 9.9 Hz, 1H), 2.36 (s, 3H); ¹³C NMR (150 MHz, CDCl₃) δ 160.3, 147.8, 137.8, 137.6, 136.0, 129.0, 127.3, 126.7 (q, *J*_{C-F} = 285.8 Hz), 123.6, 121.6, 76.5 (q, *J*_{C-F} = 27.4 Hz), 32.1, 31.2, 21.2; ¹⁹F NMR (564 MHz, CDCl₃) δ -79.90 (s, 3F); FT-IR (thin film, KBr): ν (cm⁻¹) 3107, 2928, 1596, 1435, 727; HRMS (CI) calcd C₁₆H₁₇NOF₃ [M + H]⁺: 296.1262, found: 296.1271.



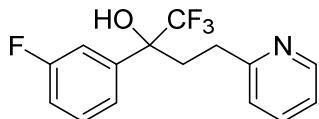
2-([1,1'-Biphenyl]-4-yl)-1,1,1-trifluoro-4-(pyridin-2-yl)butan-2-ol (3g): White solid; m.p. 188–190 °C; 94% yield (67 mg); ¹H NMR (400 MHz, CDCl₃) δ 9.45 (s, 1H), 8.51 (d, *J* = 4.3 Hz, 1H), 7.78 (d, *J* = 8.0 Hz, 2H), 7.69 – 7.54 (m, 5H), 7.44 (t, *J* = 7.4 Hz, 2H), 7.35 (t, *J* = 7.2 Hz, 1H), 7.22 – 7.10 (m, 2H), 3.00 (dd, *J* = 16.8, 8.7 Hz, 1H), 2.90 – 2.69 (m, 2H), 2.62 (dd, *J* = 14.3, 10.0 Hz, 1H); ¹³C NMR (150 MHz, CDCl₃) δ 160.2, 147.8, 140.9, 140.8, 138.1, 137.6, 128.9, 127.9, 127.5, 127.2, 127.0, 126.2 (q, *J*_{C-F} = 285.9 Hz), 123.6, 121.7, 76.5 (q, *J*_{C-F} = 27.5 Hz), 32.1, 31.2; ¹⁹F NMR (564 MHz, CDCl₃) δ -79.69 (s, 3F); FT-IR (thin film, KBr): ν (cm⁻¹) 3099, 2852, 1599, 1488, 761; HRMS (CI) calcd C₂₁H₁₉NOF₃ [M + H]⁺: 358.1419, found: 358.1430.



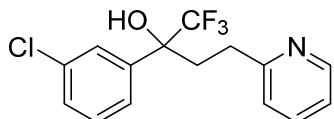
4-(1,1,1-Trifluoro-2-hydroxy-4-(pyridin-2-yl)butan-2-yl)benzonitrile (3h): White solid; m.p. 132–134 °C; 60% yield (37 mg); ¹H NMR (600 MHz, CDCl₃) δ 9.84 (s, 1H), 8.48 (d, *J* = 4.8 Hz, 1H), 7.84 (d, *J* = 8.3 Hz, 2H), 7.68 (d, *J* = 8.5 Hz, 2H), 7.65 – 7.62 (m, 1H), 7.21 (dd, *J* = 7.1, 5.4 Hz, 1H), 7.16 (d, *J* = 7.8 Hz, 1H), 3.02 – 2.93 (m, 1H), 2.70 – 2.58 (m, 3H); ¹³C NMR (150 MHz, CDCl₃) δ 159.5, 147.6, 144.7, 137.9, 132.1, 128.5, 125.8 (q, *J* = 286.1 Hz), 123.7, 121.9, 118.75, 112.1, 76.4 (q, *J* = 27.8 Hz), 32.1, 30.9; ¹⁹F NMR (564 MHz, CDCl₃) δ -79.38 (s, 3F). FT-IR (thin film, KBr): ν (cm⁻¹) 3073, 2921, 2232, 1600, 669; HRMS(CI) calcd C₁₆H₁₄F₃N₂O [M + H]⁺: 307.1058, found: 307.1059.



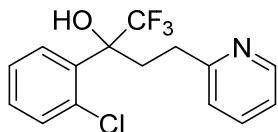
Methyl-4-(1,1,1-trifluoro-2-hydroxy-4-(pyridin-2-yl)butan-2-yl)benzoate (3i): White solid; m.p. 137–138 °C; 55% yield (37 mg); ¹H NMR (400 MHz, CDCl₃) δ 9.64 (s, 1H), 8.49 (s, 1H), 8.05 (d, *J* = 7.8 Hz, 2H), 7.79 (d, *J* = 7.6 Hz, 2H), 7.63 (t, *J* = 7.1 Hz, 1H), 7.23 – 7.07 (m, 2H), 3.91 (s, 3H), 3.11 – 2.82 (m, 1H), 2.79 – 2.45 (m, 3H); ¹³C NMR (150 MHz, CDCl₃) δ 167.0, 159.9, 147.8, 144.4, 137.7, 130.0, 129.5, 127.6, 126.1 (q, *J*_{C-F} = 286.0 Hz), 123.7, 121.8, 76.6 (q, *J*_{C-F} = 27.6 Hz), 52.3, 32.2, 31.1; ¹⁹F NMR (564 MHz, CDCl₃) δ -79.47 (s, 3F); FT-IR (thin film, KBr): ν (cm⁻¹) 3100, 1712, 1598, 1435, 713; HRMS (CI) calcd C₁₇H₁₇NO₃F₃ [M + H]⁺: 340.1161, found: 340.1163.



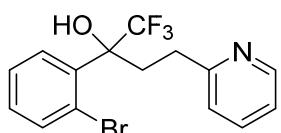
1,1,1-Trifluoro-2-(3-fluorophenyl)-4-(pyridin-2-yl)butan-2-ol (3j): White solid; m.p. 108–111 °C; 95% yield (57 mg); ¹H NMR (400 MHz, CDCl₃) δ 8.48 (s, 1H), 7.64 (t, *J* = 6.9 Hz, 1H), 7.53 – 7.39 (m, 2H), 7.40 – 7.29 (m, 1H), 7.24 – 7.08 (m, 2H), 7.07 – 6.95 (m, 1H), 2.96 (dd, *J* = 16.3, 7.8 Hz, 1H), 2.82 – 2.50 (m, 3H); ¹³C NMR (150 MHz, CDCl₃) δ 163.0 (d, *J*_{C-F} = 245.2 Hz), 159.9, 147.6, 142.0 (d, *J*_{C-F} = 6.9 Hz), 137.9, 129.7 (d, *J*_{C-F} = 8.0 Hz), 126.1 (q, *J*_{C-F} = 285.9 Hz), 123.7, 122.9, 121.8, 115.1 (d, *J*_{C-F} = 20.9 Hz), 114.9 (d, *J*_{C-F} = 20.3 Hz), 76.3 (d, *J*_{C-F} = 27.9 Hz), 32.3, 31.0; ¹⁹F NMR (564 MHz, CDCl₃) δ -79.67 (s, 3F), -112.96 – -113.07 (m, 1F); FT-IR (thin film, KBr): ν(cm⁻¹) 3064, 1482, 1477, 788, 705; HRMS (CI) calcd C₁₅H₁₄NOF₄ [M + H]⁺: 300.1012, found: 300.1021.



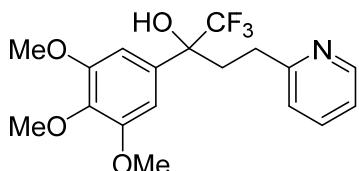
2-(3-Chlorophenyl)-1,1,1-trifluoro-4-(pyridin-2-yl)butan-2-ol (3k): White solid; m.p. 154–155 °C; 95% yield (60 mg); ¹H NMR (400 MHz, CDCl₃) δ 8.49 (d, *J* = 3.8 Hz, 1H), 7.74 (s, 1H), 7.65 (t, *J* = 7.4 Hz, 1H), 7.56 (d, *J* = 4.9 Hz, 1H), 7.38 – 7.27 (m, 2H), 7.24 – 7.09 (m, 2H), 2.97 (dd, *J* = 16.7, 8.2 Hz, 1H), 2.81 – 2.48 (m, 3H); ¹³C NMR (150 MHz, CDCl₃) δ 159.8, 147.5, 141.4, 138.0, 134.5, 129.5, 128.4, 127.9, 126.1 (q, *J*_{C-F} = 285.9 Hz), 125.5, 123.8, 121.9, 76.3 (q, *J*_{C-F} = 27.7 Hz), 32.2, 31.0; ¹⁹F NMR (564 MHz, CDCl₃) δ -79.64 (s, 3F); FT-IR (thin film, KBr): ν(cm⁻¹) 3078, 1598, 1478, 790, 713; HRMS (CI) calcd C₁₅H₁₄NOF₃³⁵Cl [M + H]⁺: 316.0716, found: 316.0726.



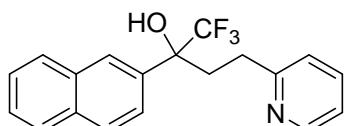
2-(2-Chlorophenyl)-1,1,1-trifluoro-4-(pyridin-2-yl)butan-2-ol (3l): Yellow solid; m.p. 117–118 °C; 33% (21 mg); ¹H NMR (600 MHz, CDCl₃) δ 9.82 (s, 1H), 8.47 (d, *J* = 4.7 Hz, 1H), 8.28 (d, *J* = 8.0 Hz, 1H), 7.63 (t, *J* = 7.7 Hz, 1H), 7.41 (d, *J* = 7.8 Hz, 1H), 7.33 (t, *J* = 7.6 Hz, 1H), 7.30 – 7.26 (m, 1H), 7.21 – 7.15 (m, 2H), 3.66 (dd, *J* = 15.3, 8.8 Hz, 1H), 3.20 (dd, *J* = 17.3, 8.8 Hz, 1H), 2.83 (dd, *J* = 17.3, 10.1 Hz, 1H), 2.53 (dd, *J* = 15.3, 10.1 Hz, 1H); ¹³C NMR (150 MHz, CDCl₃) δ 160.1, 147.6, 137.7, 136.0, 133.0, 132.3, 131.6, 129.9, 127.0, 126.2 (q, *J*_{C-F} = 286.1 Hz), 123.6, 121.7, 77.9 (q, *J*_{C-F} = 28.4 Hz), 31.9, 30.5; ¹⁹F NMR (564 MHz, CDCl₃) δ -79.11 (s, 3F); FT-IR (thin film, KBr): ν(cm⁻¹) 2920, 1597, 1459, 764; HRMS(CI) calcd C₁₅H₁₄F₃NO³⁵Cl [M + H]⁺: 316.0716, found: 316.0713.



2-(2-Bromophenyl)-1,1,1-trifluoro-4-(pyridin-2-yl)butan-2-ol (3m): White solid; m.p. 99–100 °C; 49% (35 mg); ^1H NMR (400 MHz, CDCl_3) δ 9.82 (s, 1H), 8.45 (d, J = 4.4 Hz, 1H), 8.27 (d, J = 7.8 Hz, 1H), 7.71 – 7.52 (m, 2H), 7.36 (t, J = 7.6 Hz, 1H), 7.24 – 7.06 (m, 3H), 3.79 (dd, J = 15.0, 8.9 Hz, 1H), 3.19 (dd, J = 17.2, 8.8 Hz, 1H), 2.82 (dd, J = 17.2, 10.0 Hz, 1H), 2.47 (dd, J = 15.2, 10.1 Hz, 1H); ^{13}C NMR (150 MHz, CDCl_3 , overlapping peaks) δ 160.1, 147.5, 137.7, 135.5, 133.3, 130.0, 127.5, 126.2 (q, J_{C-F} = 286.5 Hz), 123.6, 121.7, 121.3, 78.0 (q, J_{C-F} = 28.6 Hz), 31.9, 30.4; ^{19}F NMR (564 MHz, CDCl_3) δ -78.49 (s, 3F); FT-IR (thin film, KBr): ν (cm^{-1}) 2923, 2853, 1602, 1437, 760; HRMS (CI) calcd $\text{C}_{15}\text{H}_{14}\text{NOF}_3^{79}\text{Br}$ [M + H] $^+$: 360.0211, found: 360.0220.

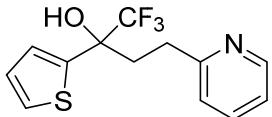


1,1,1-Trifluoro-4-(pyridin-2-yl)-2-(3,4,5-trimethoxyphenyl)butan-2-ol (3n): oil; 92% yield (68 mg); ^1H NMR (600 MHz, CDCl_3) δ 9.42 (s, 1H), 8.48 (d, J = 4.9 Hz, 1H), 7.65 – 7.59 (m, 1H), 7.19 – 7.15 (m, 1H), 7.14 (d, J = 7.8 Hz, 1H), 6.92 (s, 2H), 3.86 (s, 6H), 3.84 (s, 3H), 2.95 (dd, J = 16.8, 8.6 Hz, 1H), 2.77 (dd, J = 16.8, 9.4 Hz, 1H), 2.63 (dd, J = 14.5, 8.7 Hz, 1H), 2.54 (dd, J = 14.7, 9.5 Hz, 1H); ^{13}C NMR (150 MHz, CDCl_3) δ 160.2, 153.0, 147.7, 137.8, 137.7, 134.7, 126.3 (q, J_{C-F} = 285.8 Hz), 123.7, 121.7, 104.7, 76.5 (q, J_{C-F} = 27.4 Hz), 60.9, 56.3, 32.4, 31.1; ^{19}F NMR (564 MHz, CDCl_3) δ -79.53 (s, 3F); FT-IR (thin film, KBr): ν (cm^{-1}) 2939, 1591, 1455, 756, 711; HRMS (CI) calcd $\text{C}_{18}\text{H}_{21}\text{NO}_4\text{F}_3$ [M + H] $^+$: 372.1423, found: 372.1422.

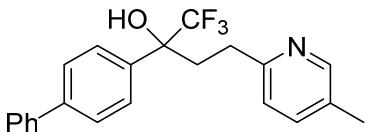


1,1,1-Trifluoro-2-(naphthalen-2-yl)-4-(pyridin-2-yl)butan-2-ol (3o): White solid; m.p. 130–131 °C; 82% yield (54 mg); ^1H NMR (400 MHz, CDCl_3) δ 8.53 (d, J = 4.1 Hz, 1H), 8.28 (s, 1H), 7.95 – 7.81 (m, 3H), 7.73 (d, J = 8.5 Hz, 1H), 7.63 (t, J = 7.4 Hz, 1H), 7.55 – 7.41 (m, 2H), 7.23 – 7.15 (m, 1H), 7.12 (d, J = 7.7 Hz, 1H), 2.96 (dd, J = 16.4, 8.5 Hz, 1H), 2.88 – 2.73 (m, 2H), 2.67 (dd, J = 13.7, 9.9 Hz, 1H); ^{13}C NMR (150 MHz, CDCl_3) δ 160.1, 147.6, 138.0, 136.4, 133.2, 133.1, 128.7, 127.9, 127.6, 127.4, 126.5, 126.4 (q, J_{C-F} = 286.1 Hz), 126.2, 124.6, 123.8, 121.8, 76.7 (q, J_{C-F} = 27.5 Hz), 32.3, 31.1; ^{19}F NMR (564 MHz, CDCl_3) δ -79.43 (s, 3F); FT-IR (thin film,

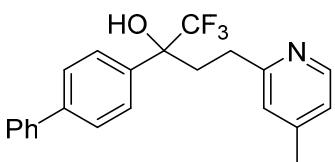
KBr): ν (cm⁻¹) 3064, 2923, 1595, 1459, 753; HRMS (CI) calcd C₁₉H₁₇NOF₃ [M + H]⁺: 332.1262, found: 332.1272.



1,1,1-Trifluoro-4-(pyridin-2-yl)-2-(thiophen-2-yl)butan-2-ol (3p): Yellow oil; 33% yield (19 mg); ¹H NMR (600 MHz, CDCl₃) δ 10.15 (s, 1H), 8.48 (d, *J* = 4.9 Hz, 1H), 7.64 (t, *J* = 7.7 Hz, 1H), 7.30 (d, *J* = 5.0 Hz, 1H), 7.22 – 7.14 (m, 2H), 7.11 (d, *J* = 2.9 Hz, 1H), 7.07 – 7.02 (m, 1H), 3.06 – 2.97 (m, 1H), 2.96 – 2.86 (m, 1H), 2.72 – 2.31 (m, 2H); ¹³C NMR (150 MHz, CDCl₃) δ 160.0, 147.7, 144.9, 137.8, 127.4, 126.0, 125.7 (q, *J*_{C-F} = 285.6 Hz), 125.1, 123.7, 121.8, 76.2 (q, *J*_{C-F} = 29.3 Hz), 33.6, 31.2; ¹⁹F NMR (564 MHz, CDCl₃) δ -81.00 (s, 3F); FT-IR (thin film, KBr): ν (cm⁻¹) 3082, 2920, 1597, 1437, 707; HRMS (CI) calcd C₁₃H₁₃F₃NOS [M + H]⁺: 288.0670, found: 288.0668.

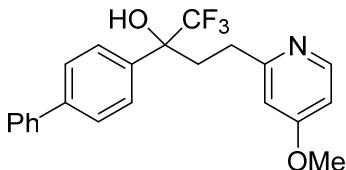


2-([1,1'-Biphenyl]-4-yl)-1,1,1-trifluoro-4-(5-methylpyridin-2-yl)butan-2-ol (3q): White solid; m.p. 167-169 °C; 60% yield (44 mg); ¹H NMR (600 MHz, CDCl₃) δ 9.61 (s, 1H), 8.33 (s, 1H), 7.78 (d, *J* = 8.1 Hz, 2H), 7.62 (t, *J* = 7.5 Hz, 4H), 7.48 – 7.39 (m, 3H), 7.35 (t, *J* = 7.3 Hz, 1H), 7.03 (d, *J* = 7.9 Hz, 1H), 2.95 (dd, *J* = 16.8, 8.9 Hz, 1H), 2.81 – 2.67 (m, 2H), 2.59 (dd, *J* = 14.9, 9.9 Hz, 1H), 2.31 (s, 3H); ¹³C NMR (150 MHz, CDCl₃) δ 157.1, 147.9, 140.9, 140.8, 138.3, 138.2, 131.2, 128.9, 127.9, 127.5, 127.2, 126.9, 126.4 (q, *J*_{C-F} = 285.9 Hz), 123.1, 76.5 (q, *J*_{C-F} = 27.4 Hz), 32.3, 30.7, 18.1; ¹⁹F NMR (564 MHz, CDCl₃) δ -79.68 (s, 3F); FT-IR (thin film, KBr): ν (cm⁻¹) 3100, 2857, 1609, 1496, 763; HRMS (CI) calcd C₂₂H₂₁F₃NO [M + H]⁺: 372.1575, found: 372.1571.

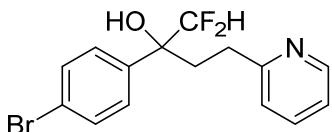


2-([1,1'-Biphenyl]-4-yl)-1,1,1-trifluoro-4-(4-methylpyridin-2-yl)butan-2-ol (3r): White solid; m.p. 130-131 °C; 57% yield (42 mg); ¹H NMR (600 MHz, CDCl₃) δ 8.35 (d, *J* = 5.2 Hz, 1H), 7.78 (d, *J* = 8.1 Hz, 2H), 7.62 (t, *J* = 7.5 Hz, 4H), 7.44 (t, *J* = 7.6 Hz, 2H), 7.35 (t, *J* = 7.3 Hz, 1H), 7.00 (d, *J* = 5.1 Hz, 1H), 6.96 (s, 1H), 2.92 (dd, *J* = 16.8, 8.8 Hz, 1H), 2.82 – 2.66 (m, 2H), 2.60 (dd, *J* = 14.8, 9.7 Hz, 1H), 2.31 (s, 3H); ¹³C NMR (150 MHz, CDCl₃) δ 159.8, 149.2, 147.3, 140.9, 140.7, 138.2, 128.9, 127.9, 127.5, 127.2, 127.0, 126.4 (q, *J*_{C-F} = 286.0 Hz), 124.4, 122.7, 76.5 (q, *J*_{C-F} =

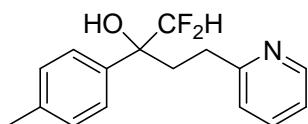
27.5 Hz), 32.2, 31.0, 21.2; ^{19}F NMR (564 MHz, CDCl_3) δ -79.66 (s, 3F); FT-IR (thin film, KBr): ν (cm^{-1}) 3065, 2923, 1612, 1487, 732; HRMS (CI) calcd $\text{C}_{22}\text{H}_{21}\text{F}_3\text{NO}$ [$\text{M} + \text{H}]^+$: 372.1575, found: 375.1580.



2-([1,1'-Biphenyl]-4-yl)-1,1,1-trifluoro-4-(4-methoxypyridin-2-yl)butan-2-ol (3s): White solid; m.p. 107-109 °C; 72% yield (56 mg); ^1H NMR (600 MHz, CDCl_3) δ 8.32 (d, $J = 5.9$ Hz, 1H), 7.79 (d, $J = 8.1$ Hz, 2H), 7.62 (t, $J = 8.1$ Hz, 4H), 7.44 (t, $J = 7.5$ Hz, 2H), 7.35 (t, $J = 7.3$ Hz, 1H), 6.70 (d, $J = 5.5$ Hz, 1H), 6.63 (s, 1H), 3.81 (s, 3H), 2.90 (dd, $J = 16.7, 8.8$ Hz, 1H), 2.81 – 2.67 (m, 2H), 2.60 (dd, $J = 14.7, 9.9$ Hz, 1H); ^{13}C NMR (150 MHz, CDCl_3) δ 166.7, 161.8, 148.9, 140.8, 140.7, 138.2, 128.9, 127.9, 127.5, 127.2, 126.9, 126.4 (q, $J_{\text{C}-\text{F}} = 285.8$ Hz), 109.1, 108.1, 76.4 (q, $J_{\text{C}-\text{F}} = 27.6$ Hz), 55.3, 32.1, 31.4; ^{19}F NMR (564 MHz, CDCl_3) δ -79.64 (s, 3F); FT-IR (thin film, KBr): ν (cm^{-1}) 2922, 2851, 1609, 1486, 692; HRMS(CI) calcd $\text{C}_{22}\text{H}_{21}\text{F}_3\text{NO}_2$ [$\text{M} + \text{H}]^+$: 388.1524, found: 388.1523.

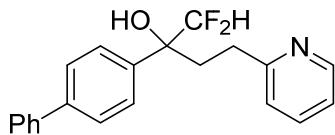


2-(4-Bromophenyl)-1,1-difluoro-4-(pyridin-2-yl)butan-2-ol (5a): White solid; m.p. 112-113 °C; 63% yield (43mg); ^1H NMR (600 MHz, CDCl_3) δ 8.87 (s, 1H), 8.48 (d, $J = 4.8$ Hz, 1H), 7.62 (t, $J = 7.7$ Hz, 1H), 7.52 (d, $J = 8.7$ Hz, 2H), 7.50 (d, $J = 8.7$ Hz, 2H), 7.19 – 7.15 (m, 1H), 7.13 (d, $J = 7.8$ Hz, 1H), 5.70 (t, $J = 56.5$ Hz, 1H), 2.95 (dd, $J = 16.8, 8.6$ Hz, 1H), 2.74 (dd, $J = 16.7, 9.7$ Hz, 1H), 2.55 (dd, $J = 14.8, 8.6$ Hz, 1H), 2.43 (dd, $J = 14.9, 9.8$ Hz, 1H); ^{13}C NMR (150 MHz, CDCl_3) δ 160.3, 147.8, 139.7, 137.6, 131.4, 129.2, 123.7, 122.0, 121.7, 117.6 (t, $J_{\text{C}-\text{F}} = 250.5$ Hz), 75.6 (t, $J_{\text{C}-\text{F}} = 20.8$ Hz), 31.4, 31.3; ^{19}F NMR (564 MHz, CDCl_3) δ -128.22 (dd, $J = 273.5, 56.8$ Hz, 1F), -129.52 (dd, $J = 273.5, 56.3$ Hz, 1F); FT-IR (thin film, KBr): ν (cm^{-1}) 3184, 2925, 1596, 1434, 809; HRMS (CI) calcd $\text{C}_{15}\text{H}_{15}^{79}\text{BrF}_2\text{NO}$ [$\text{M} + \text{H}]^+$: 342.0305, found: 342.0310.

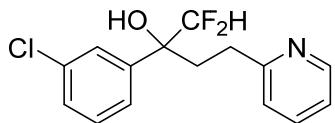


1,1-Difluoro-4-(pyridin-2-yl)-2-(p-tolyl)butan-2-ol (5b): White solid; m.p. 114-116°C; 60% yield (33 mg); ^1H NMR (400 MHz, CDCl_3) δ 8.48 (d, $J = 3.8$ Hz, 1H), 7.60 (t, $J = 7.4$ Hz, 1H), 7.52 (d, $J = 7.8$ Hz, 2H), 7.20 (d, $J = 7.8$ Hz, 2H), 7.18 –

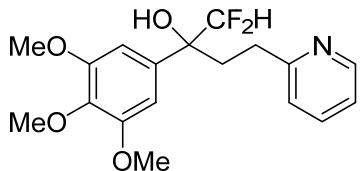
7.03 (m, 2H), 5.72 (t, $J = 56.6$ Hz, 1H), 3.00 – 2.86 (m, 1H), 2.88 – 2.72 (m, 1H), 2.65 – 2.52 (m, 1H), 2.50 – 2.38 (m, 1H), 2.35 (s, 3H); ^{13}C NMR (150 MHz, CDCl_3) δ 160.7, 147.8, 137.5, 137.4, 137.3, 129.0, 127.1, 123.6, 121.5, 117.9 (t, $J_{\text{C}-\text{F}} = 250.1$ Hz), 75.7 (t, $J_{\text{C}-\text{F}} = 20.5$ Hz), 31.5, 31.3, 21.1; ^{19}F NMR (564 MHz, CDCl_3) δ -128.66 (dd, $J = 272.1, 56.9$ Hz, 1F), -129.65 (dd, $J = 272.1, 56.3$ Hz, 1F); FT-IR (thin film, KBr): ν (cm^{-1}) 3180, 2926, 1593, 1434, 743; HRMS (CI) calcd $\text{C}_{16}\text{H}_{18}\text{NOF}_2$ [M + H] $^+$: 278.1356, found: 278.1368.



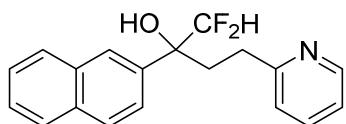
2-([1,1'-Biphenyl]-4-yl)-1,1-difluoro-4-(pyridin-2-yl)butan-2-ol (5c): White solid; m.p. 159–161 °C; 62% yield (42 mg); ^1H NMR (400 MHz, CDCl_3) δ 8.77 (s, 1H), 8.51 (s, 1H), 7.72 (d, $J = 7.5$ Hz, 2H), 7.68 – 7.54 (m, 5H), 7.48 – 7.38 (m, 2H), 7.38 – 7.29 (m, 1H), 7.18 – 7.08 (m, 2H), 5.79 (t, $J = 56.6$ Hz, 1H), 3.09 – 2.92 (m, 1H), 2.93 – 2.74 (m, 1H), 2.75 – 2.56 (m, 1H), 2.56 – 2.35 (m, 1H); ^{13}C NMR (150 MHz, CDCl_3) δ 160.6, 147.9, 140.9, 140.5, 139.6, 137.4, 128.9, 127.7, 127.4, 127.2, 127.0, 123.6, 121.5, 117.8 (t, $J_{\text{C}-\text{F}} = 250.3$ Hz), 75.8 (t, $J_{\text{C}-\text{F}} = 20.6$ Hz), 31.5, 31.4; ^{19}F NMR (564 MHz, CDCl_3) δ -128.40 (dd, $J = 272.5, 56.9$ Hz, 1F), -129.45 (dd, $J = 272.5, 56.3$ Hz, 1F); FT-IR (thin film, KBr): ν (cm^{-1}) 3120, 2874, 1596, 1437, 759; HRMS (CI) calcd $\text{C}_{21}\text{H}_{20}\text{NOF}_2$ [M + H] $^+$: 340.1513, found: 340.1519.



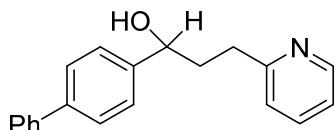
2-(3-Chlorophenyl)-1,1-difluoro-4-(pyridin-2-yl)butan-2-ol (5d): oil; 72% yield (43 mg); ^1H NMR (600 MHz, CDCl_3) δ 8.95 (s, 1H), 8.48 (d, $J = 4.8$ Hz, 1H), 7.68 (s, 1H), 7.62 (t, $J = 7.7$ Hz, 1H), 7.50 (d, $J = 7.5$ Hz, 1H), 7.33 – 7.26 (m, 2H), 7.19 – 7.15 (m, 1H), 7.14 (d, $J = 7.8$ Hz, 1H), 5.71 (t, $J = 56.5$ Hz, 1H), 2.96 (dd, $J = 16.7, 8.6$ Hz, 1H), 2.75 (dd, $J = 16.7, 9.6$ Hz, 1H), 2.56 (dd, $J = 14.9, 8.7$ Hz, 1H), 2.44 (dd, $J = 14.9, 9.8$ Hz, 1H); ^{13}C NMR (150 MHz, CDCl_3) δ 160.3, 147.8, 142.9, 137.6, 134.4, 129.5, 127.9, 127.7, 125.4, 123.6, 121.6, 117.6 (t, $J_{\text{C}-\text{F}} = 250.6$ Hz), 75.61 (t, $J_{\text{C}-\text{F}} = 20.9$ Hz), 31.5, 31.3; ^{19}F NMR (564 MHz, CDCl_3) δ -128.22 (dd, $J = 273.6, 56.8$ Hz, 1F), -129.46 (dd, $J = 273.6, 56.2$ Hz, 1F); FT-IR (thin film, KBr): ν (cm^{-1}) 3169, 1597, 1435, 736, 696; HRMS (CI) calcd $\text{C}_{15}\text{H}_{15}^{35}\text{ClF}_2\text{NO}$ [M + H] $^+$: 298.0810, found 298.0809.



1, 1-Difluoro-4-(pyridin-2-yl)-2-(3, 4, 5-trimethoxyphenyl) butan-2-ol (5e): oil; 65% yield (46 mg); ¹H NMR (600 MHz, CDCl₃) δ 8.77 (s, 1H), 8.49 (d, *J* = 4.9 Hz, 1H), 7.62 (t, *J* = 7.6 Hz, 1H), 7.19 – 7.16 (m, 1H), 7.14 (d, *J* = 7.8 Hz, 1H), 6.86 (s, 2H), 5.72 (t, *J* = 56.6 Hz, 1H), 3.87 (s, 6H), 3.85 (s, 3H), 2.95 (dd, *J* = 16.5, 8.1 Hz, 1H), 2.81 (dd, *J* = 16.6, 9.2 Hz, 1H), 2.54 (dd, *J* = 14.5, 8.4 Hz, 1H), 2.42 (dd, *J* = 14.7, 9.4 Hz, 1H); ¹³C NMR (150 MHz, CDCl₃) δ 160.6, 153.0, 147.9, 137.5, 137.4, 136.3, 123.67, 121.6, 117.8 (t, *J*_{C-F} = 250.3 Hz), 104.4, 75.8 (t, *J*_{C-F} = 20.6 Hz), 60.9, 56.3, 31.6, 31.3; ¹⁹F NMR (564 MHz, CDCl₃) δ -128.12 (dd, *J* = 272.0, 57.0 Hz, 1F), -129.18 (dd, *J* = 272.0, 56.3 Hz, 1F); FT-IR (thin film, KBr): ν (cm⁻¹) 2922, 1590, 1508, 759, 707; HRMS (CI) calcd C₁₈H₂₂F₂NO₄ [M + H]⁺: 354.1517, found: 354.1521.

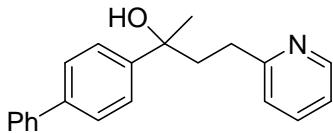


1,1-Difluoro-2-(naphthalen-2-yl)-4-(pyridin-2-yl)butan-2-ol (5f): White solid; m.p. 142–143 °C; 64% yield (40 mg); ¹H NMR (400 MHz, CDCl₃) δ 8.91 (s, 1H), 8.52 (d, *J* = 4.1 Hz, 1H), 8.22 (s, 1H), 7.95 – 7.78 (m, 3H), 7.69 (d, *J* = 8.5 Hz, 1H), 7.59 (t, *J* = 7.4 Hz, 1H), 7.48 (dd, *J* = 5.8, 3.0 Hz, 2H), 7.21 – 7.13 (m, 1H), 7.10 (d, *J* = 7.7 Hz, 1H), 5.82 (t, *J* = 56.5 Hz, 1H), 2.96 (dd, *J* = 16.5, 8.5 Hz, 1H), 2.85 – 2.66 (m, 2H), 2.52 (dd, *J* = 14.0, 10.4 Hz, 1H); ¹³C NMR (100 MHz, CDCl₃) δ 160.6, 147.9, 138.1, 137.4, 133.3, 132.9, 128.5, 127.9, 127.6, 127.0, 126.2, 126.1, 124.8, 123.6, 121.5, 118.0 (q, *J*_{C-F} = 250.7 Hz), 76.0 (t, *J*_{C-F} = 20.6 Hz), 31.4, 31.3; ¹⁹F NMR (376 MHz, CDCl₃) δ -128.08 (d, *J* = 272.9 Hz, 1F), -129.21 (d, *J* = 272.9 Hz, 1F); FT-IR (thin film, KBr): ν (cm⁻¹) 3147, 1595, 1435, 750, 670; HRMS (CI) calcd C₁₉H₁₈NOF₂ [M + H]⁺: 314.1356, found: 314.1364.

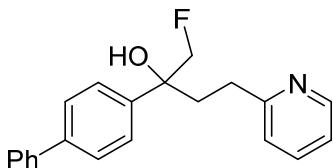


1-([1,1'-Biphenyl]-4-yl)-3-(pyridin-2-yl)propan-1-ol (7a): White solid; m.p. 88–92 °C; 60% yield (35 mg); ¹H NMR (600 MHz, CDCl₃) δ 8.52 (d, *J* = 4.4 Hz, 1H), 7.62 (t, *J* = 7.8 Hz, 1H), 7.59 (d, *J* = 7.8 Hz, 2H), 7.57 (d, *J* = 8.1 Hz, 2H), 7.47 (d, *J* = 8.1 Hz, 2H), 7.43 (t, *J* = 7.6 Hz, 2H), 7.34 (t, *J* = 7.3 Hz, 1H), 7.19 (d, *J* = 7.8 Hz, 1H), 7.17 – 7.10 (m, 1H), 4.88 (dd, *J* = 8.0, 4.1 Hz, 1H), 3.07 – 2.98 (m, 2H), 2.30 – 2.17 (m, 2H); ¹³C NMR (150 MHz, CDCl₃) δ 161.4, 148.5, 144.4, 141.1, 140.0, 137.1, 128.8, 127.22, 127.16, 127.13, 126.4, 123.5, 121.4, 73.5, 38.1, 34.5; FT-IR (thin film, KBr): ν (cm⁻¹) 3197, 2922, 1593, 1447, 759; HRMS (ESI) calcd C₂₀H₂₀NO [M + H]⁺:

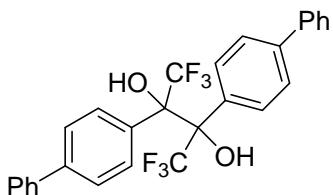
290.1539, found: 290.1539.



2-([1,1'-Biphenyl]-4-yl)-4-(pyridin-2-yl)butan-2-ol (7b): oil; 19% yield (11 mg); ^1H NMR (400 MHz, CDCl_3) δ 8.49 (d, $J = 3.3$ Hz, 1H), 7.61 (d, $J = 7.0$ Hz, 4H), 7.59 – 7.55 (m, 3H), 7.43 (t, $J = 7.6$ Hz, 2H), 7.33 (t, $J = 7.3$ Hz, 1H), 7.14 – 7.07 (m, 2H), 2.83 (t, $J = 6.6$ Hz, 2H), 2.42 – 2.36 (m, 1H), 2.33 – 2.28 (m, 1H), 1.66 (s, 3H); ^{13}C NMR (150 MHz, CDCl_3) δ 161.7, 148.3, 147.9, 141.1, 139.1, 137.0, 128.8, 127.14, 127.13, 126.8, 125.8, 123.41, 121.2, 73.8, 42.2, 32.9, 31.5; FT-IR (thin film, KBr): ν (cm^{-1}) 2960, 2923, 1596, 1457, 760; HRMS (ESI) calcd $\text{C}_{21}\text{H}_{22}\text{NO}$ [$\text{M} + \text{H}]^+$: 304.1696, found: 304.1696.



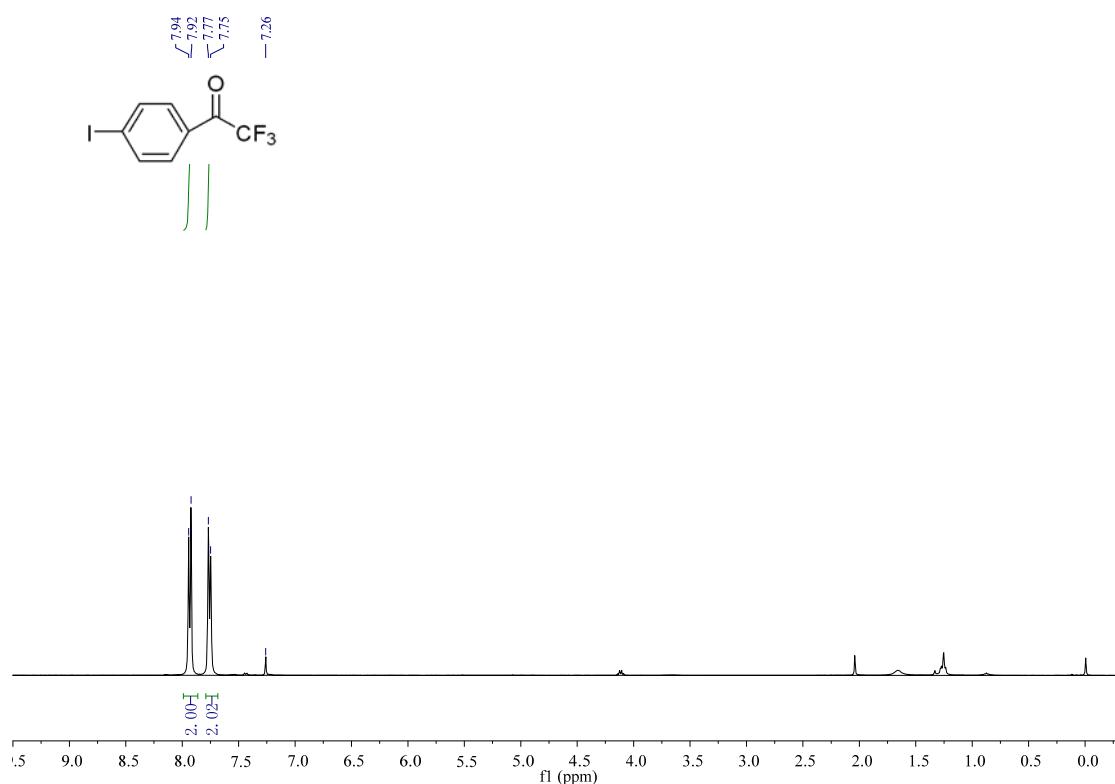
2-([1,1'-Biphenyl]-4-yl)-1-fluoro-4-(pyridin-2-yl)butan-2-ol (7c): oil; 18% yield (12 mg); ^1H NMR (600 MHz, CDCl_3) δ 8.51 (d, $J = 4.6$ Hz, 1H), 7.65 (d, $J = 8.3$ Hz, 2H), 7.62 – 7.57 (m, 5H), 7.43 (t, $J = 7.7$ Hz, 2H), 7.34 (t, $J = 7.4$ Hz, 1H), 7.16 – 7.11 (m, 2H), 4.50 (d, $J = 47.9$ Hz, 2H), 2.95 – 2.80 (m, 2H), 2.56 – 2.45 (m, 2H); ^{13}C NMR (150 MHz, CDCl_3) δ 161.2, 148.18, 142.55, 140.97, 140.04, 137.24, 128.8, 127.3, 127.2, 127.1, 126.7, 123.5, 121.4, 89.9 (d, $J_{\text{C}-\text{F}} = 179.8$ Hz), 75.1 (d, $J_{\text{C}-\text{F}} = 17.7$ Hz), 35.1, 32.0; ^{19}F NMR (564 MHz, CDCl_3) δ -222.10 (t, $J = 47.9$ Hz, 1F); FT-IR (thin film, KBr): ν (cm^{-1}) 3167, 2923, 1597, 1436, 759; HRMS (CI) calcd $\text{C}_{21}\text{H}_{21}\text{FNO}$ [$\text{M} + \text{H}]^+$: 322.1607, found: 322.1611.



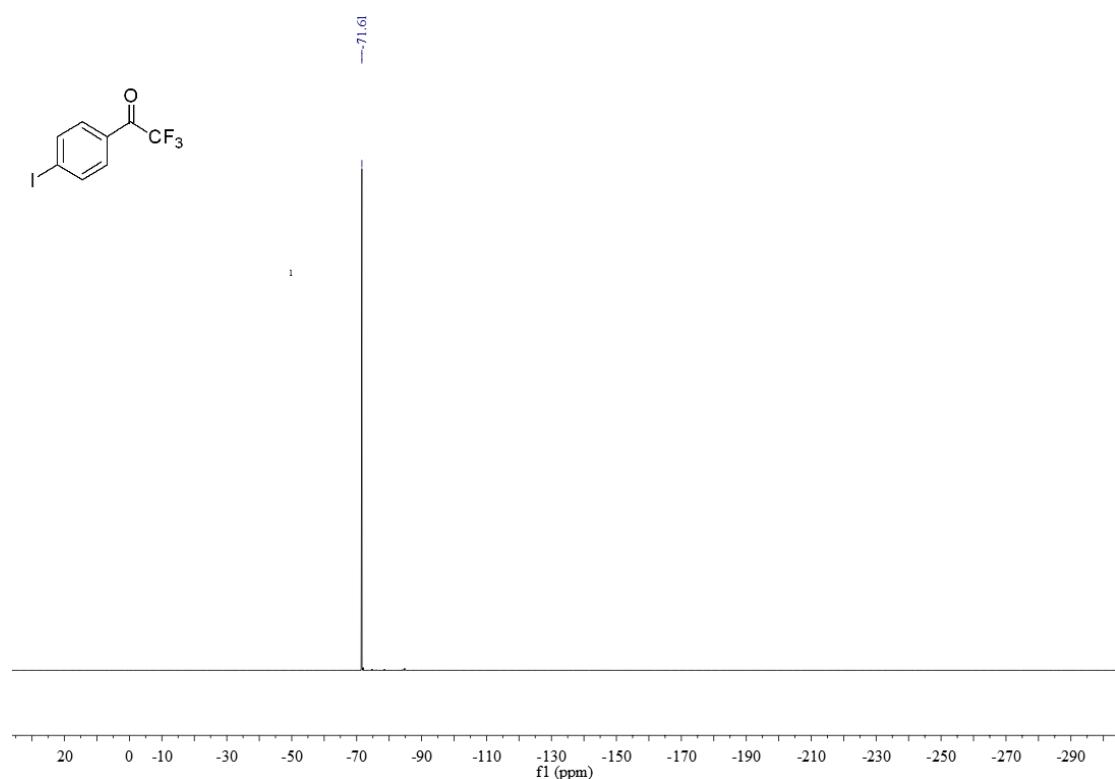
2,3-Bi([1,1'-biphenyl]-4-yl)-1,1,1,4,4,4-hexafluorobutane-2,3-diol (9): White solid; m.p. 308–310 °C; 32% yield (32 mg); ^1H NMR (600 MHz, $(\text{CD}_3)_2\text{CO}$) δ 7.91 (d, $J = 8.4$ Hz, 4H), 7.71 (d, $J = 7.4$ Hz, 4H), 7.68 (d, $J = 8.5$ Hz, 4H), 7.48 (t, $J = 7.7$ Hz, 4H), 7.38 (t, $J = 7.4$ Hz, 2H), 6.42 (s, 2H); ^{13}C NMR (150 MHz, $(\text{CD}_3)_2\text{CO}$) δ 141.8, 141.0, 135.9, 129.7, 129.3, 128.4, 127.7, 126.5, 126.1 (q, $J_{\text{C}-\text{F}} = 289.4$ Hz), 81.2 (q, $J_{\text{C}-\text{F}} = 26.4$ Hz); ^{19}F NMR (564 MHz, $(\text{CD}_3)_2\text{CO}$) δ -74.81 (s, 6F); FT-IR (thin film, KBr): ν (cm^{-1}) 3546, 1488, 1407, 721, 693; HRMS (ESI) calcd $\text{C}_{28}\text{H}_{21}\text{F}_6\text{O}_2$ [$\text{M} + \text{H}]^+$: 503.1440, found: 503.1443.

8. NMR spectra for the substrates and products

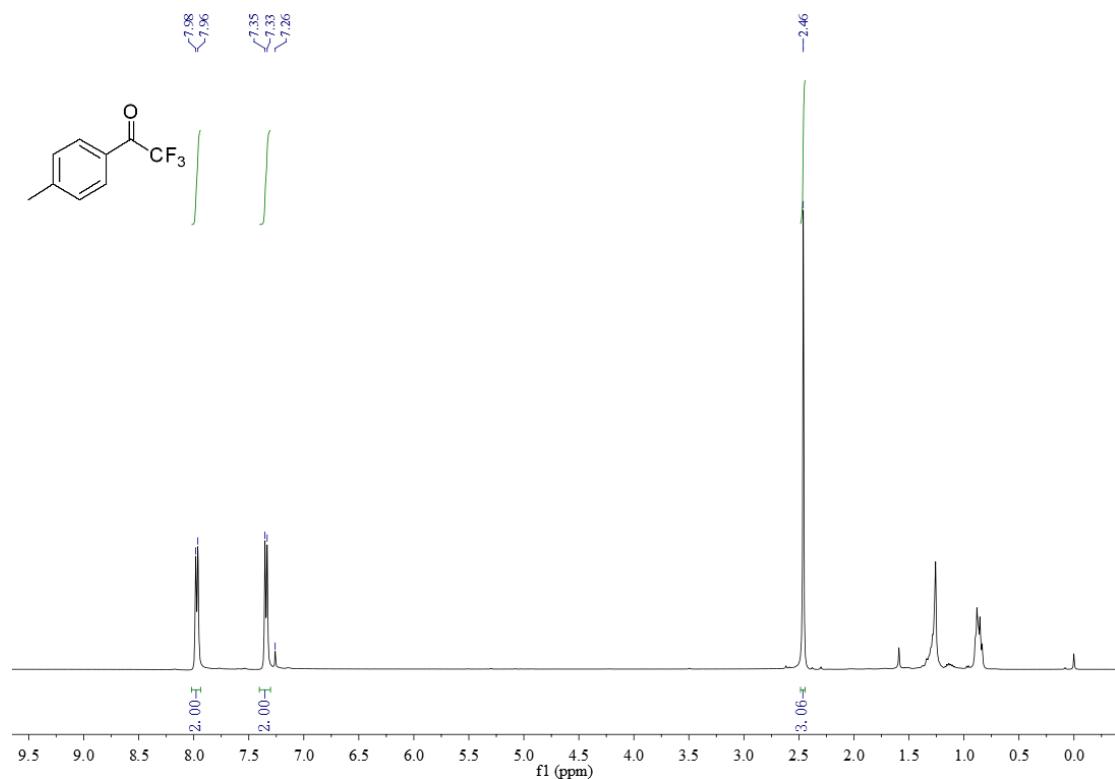
¹H NMR of **1e**



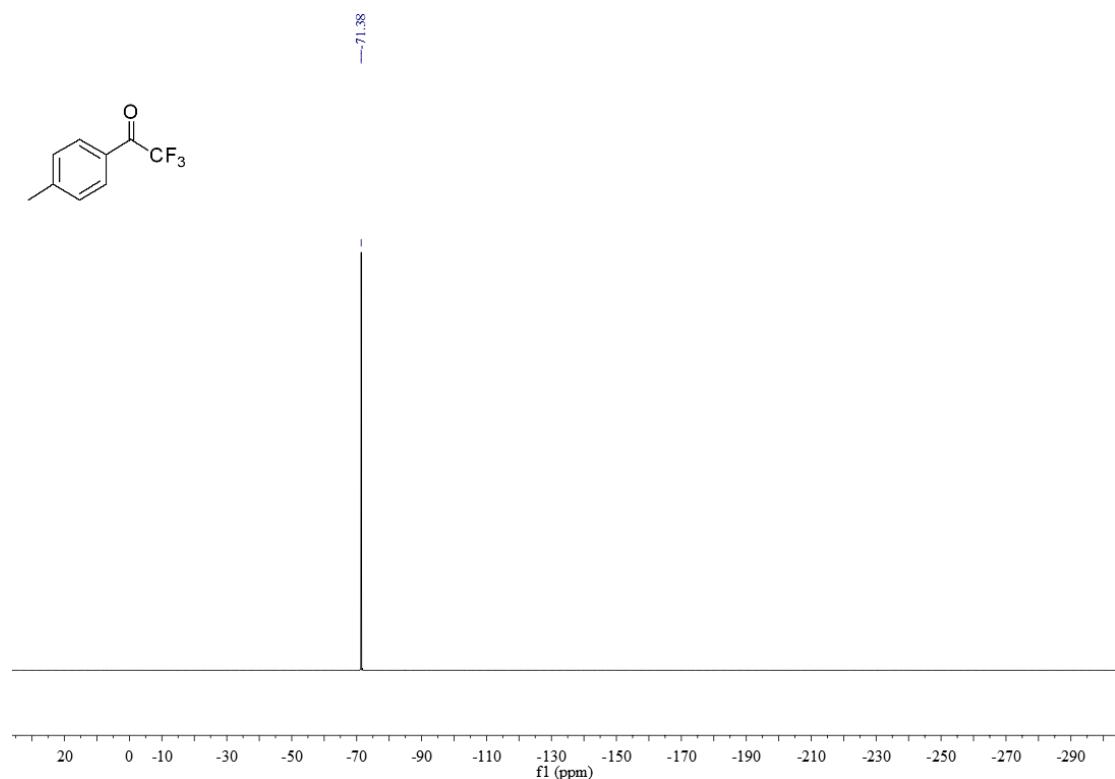
¹⁹F NMR of **1e**



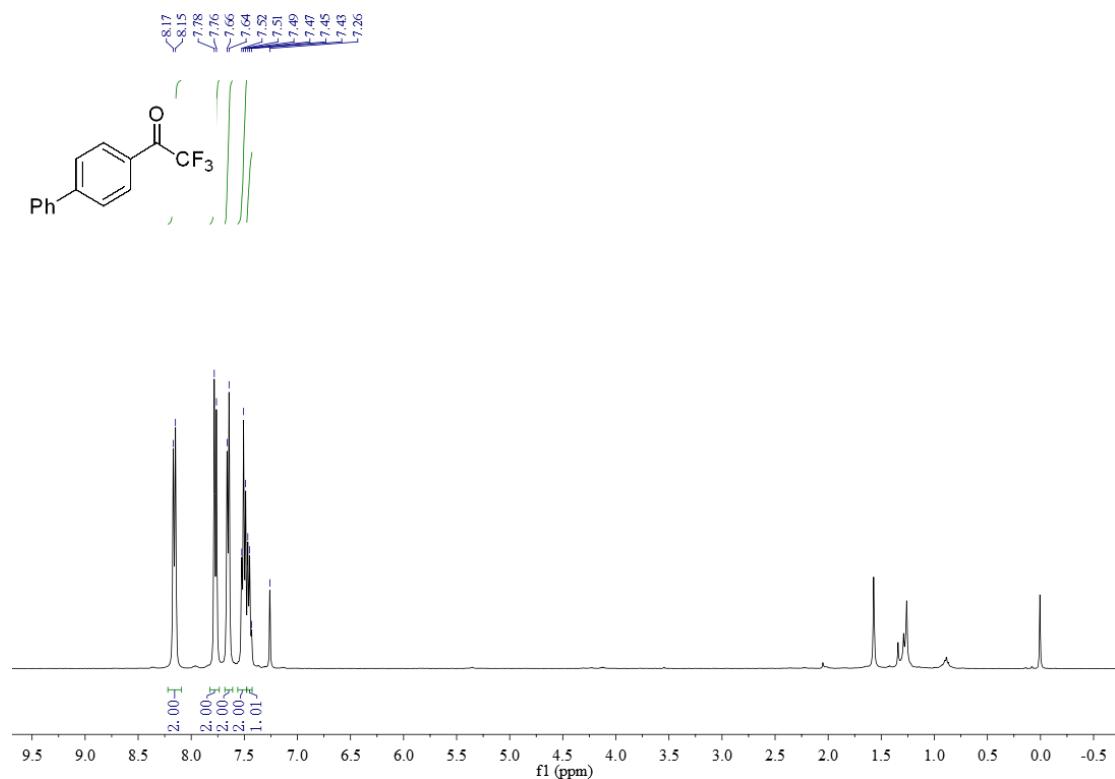
¹H NMR of **1f**



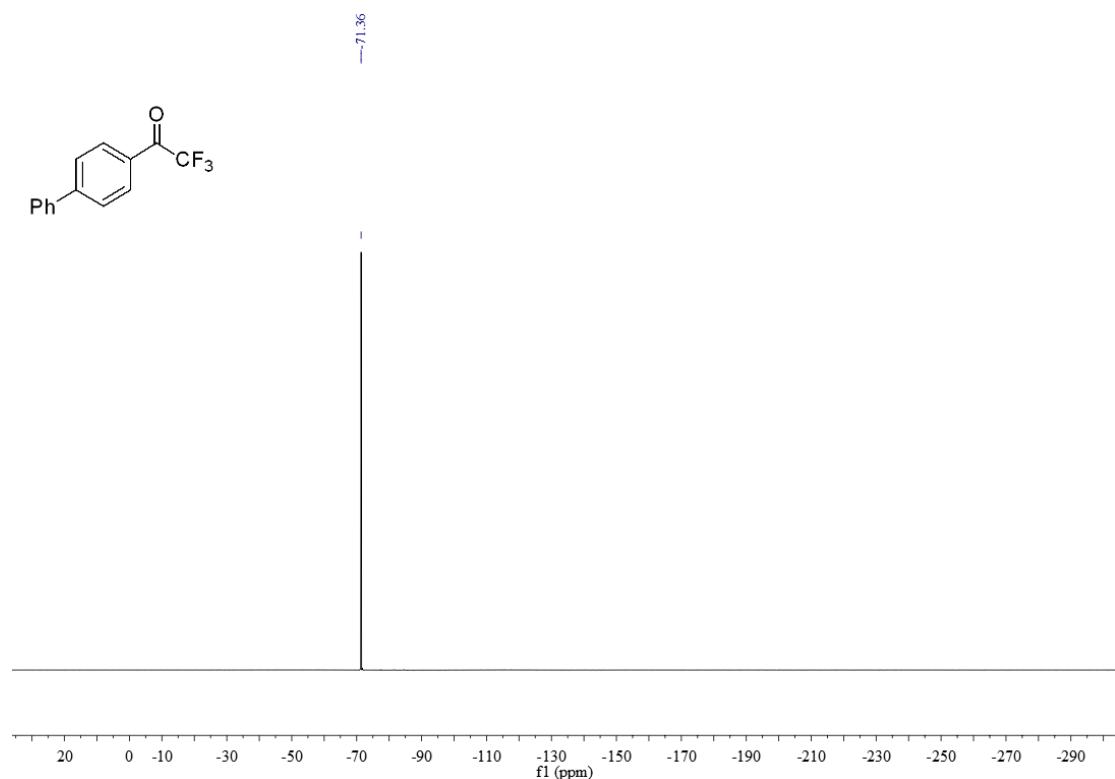
¹⁹F NMR of **1f**



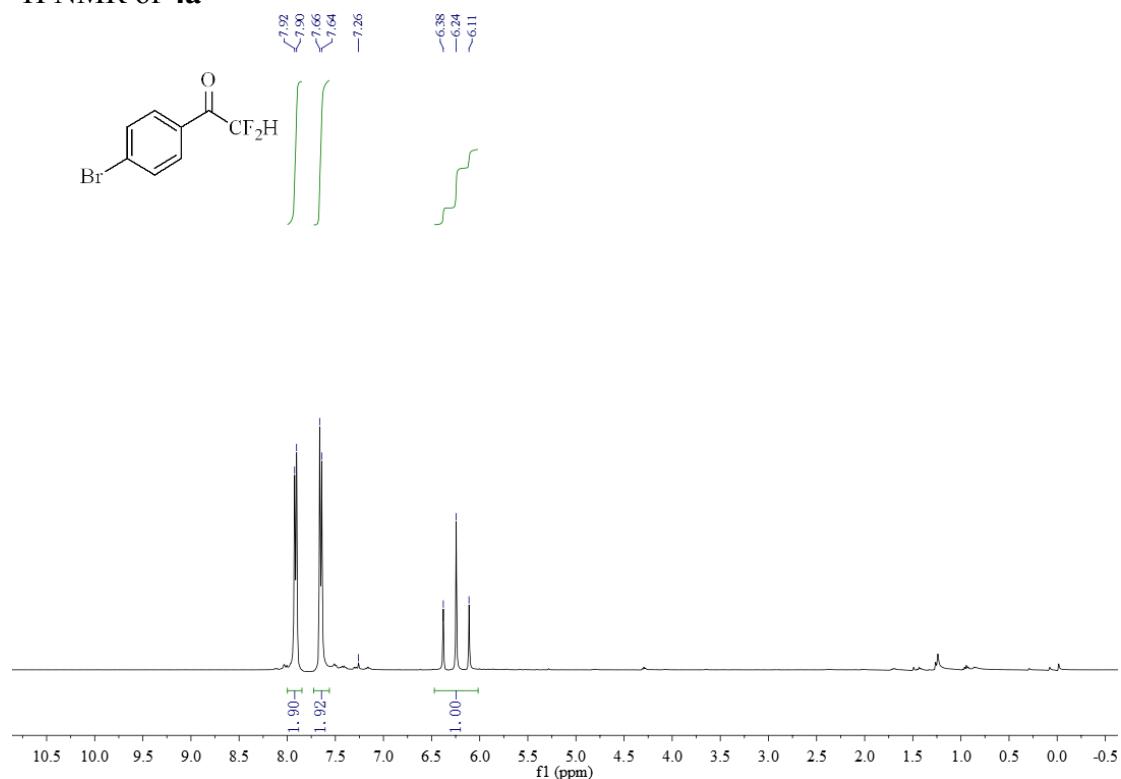
¹H NMR of **1g**



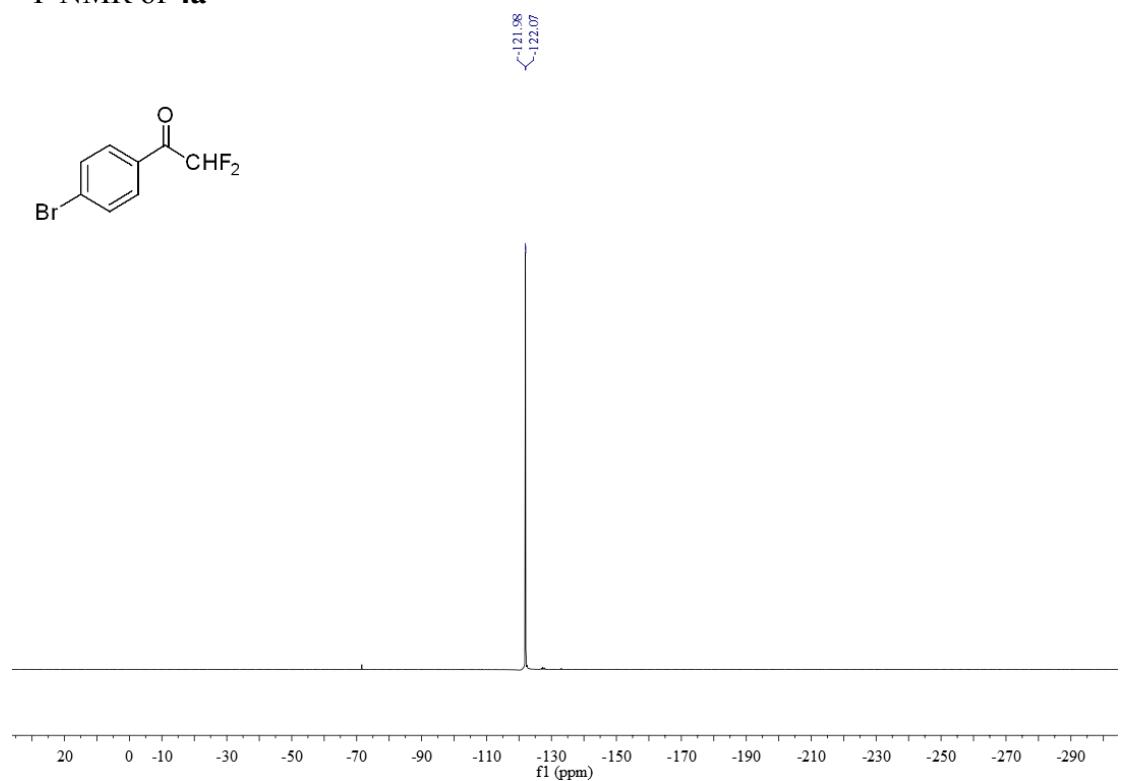
¹⁹F NMR of **1g**



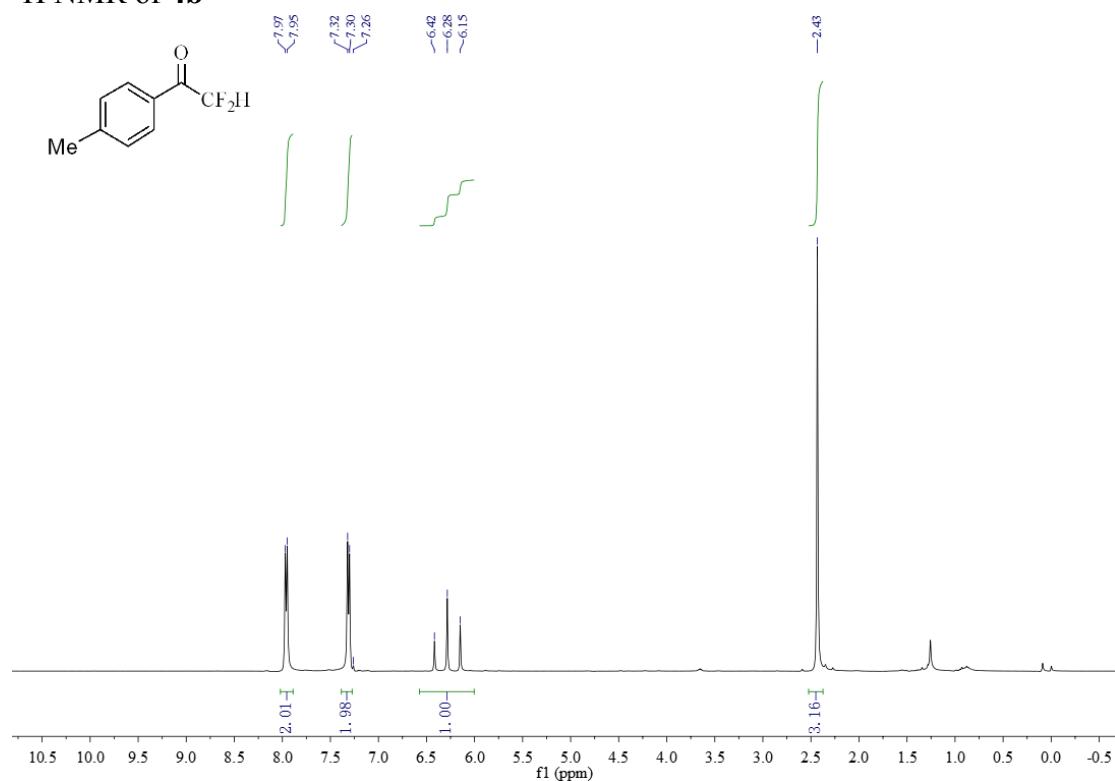
¹H NMR of **4a**



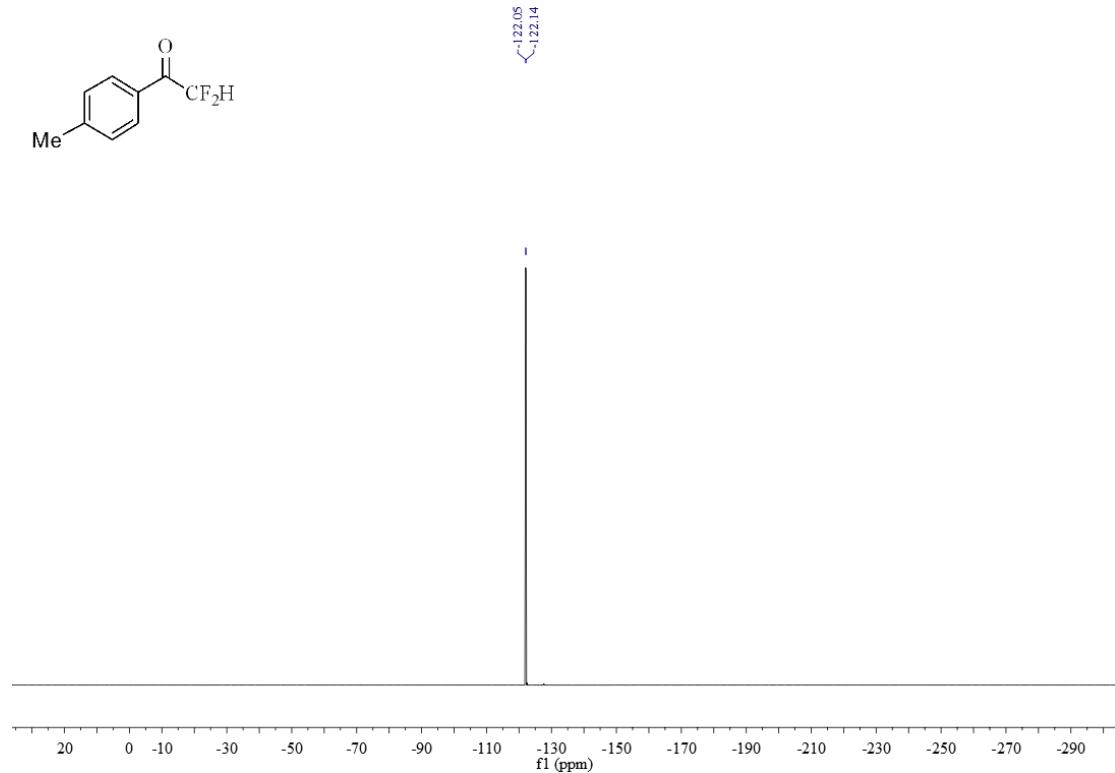
¹⁹F NMR of **4a**



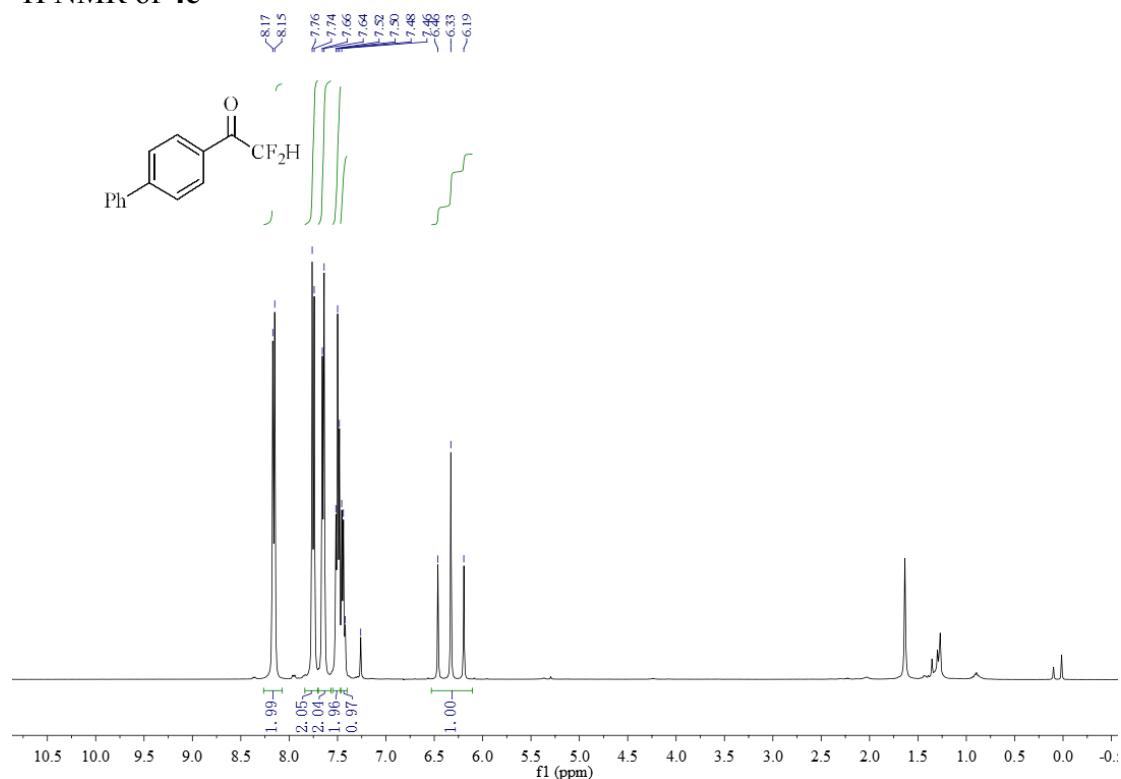
¹H NMR of **4b**



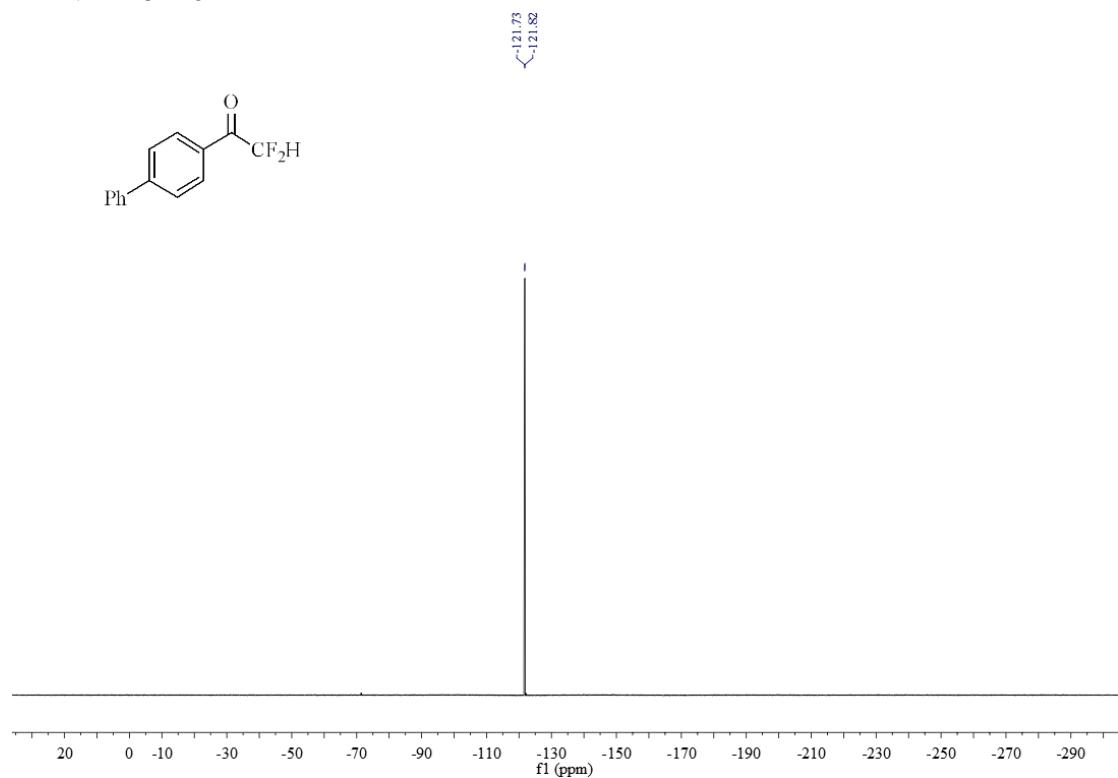
¹⁹F NMR of **4b**



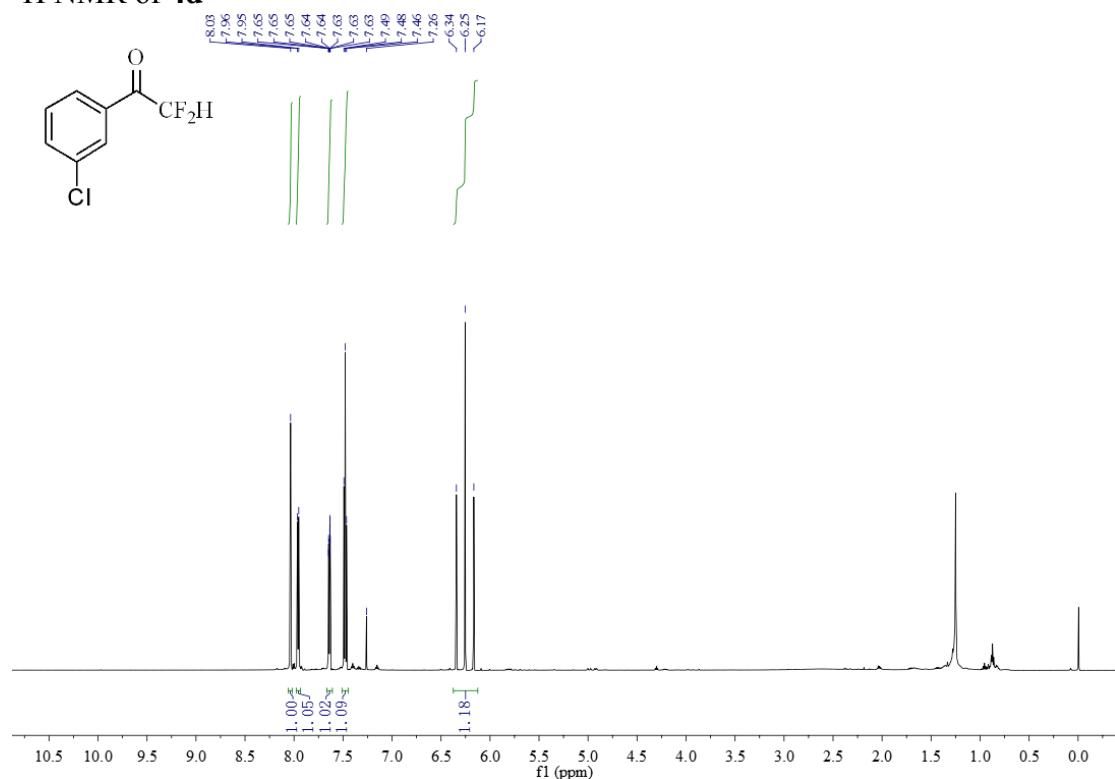
¹H NMR of **4c**



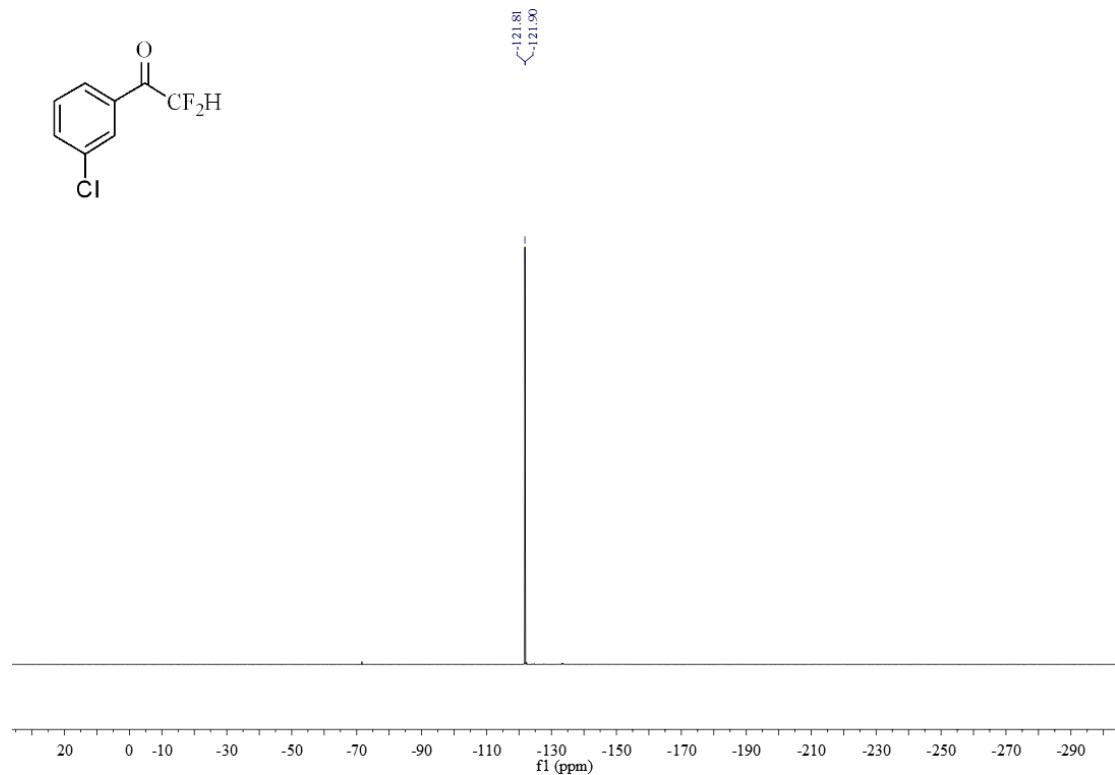
¹⁹F NMR of **4c**



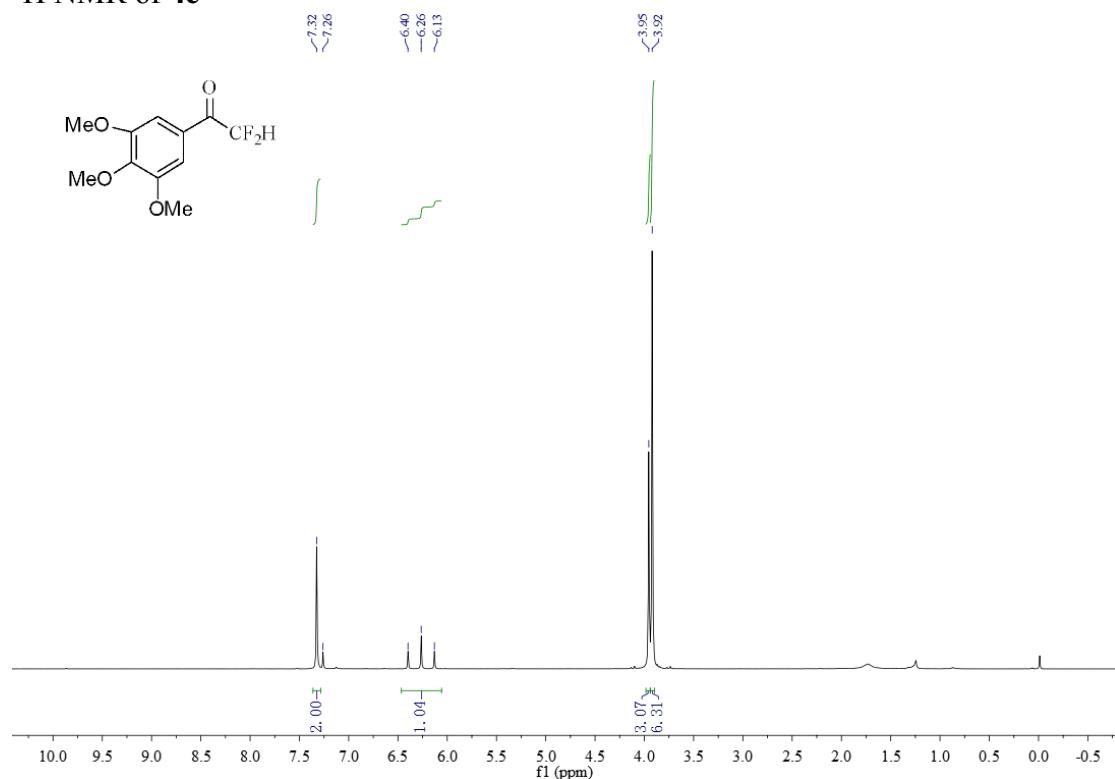
¹H NMR of **4d**



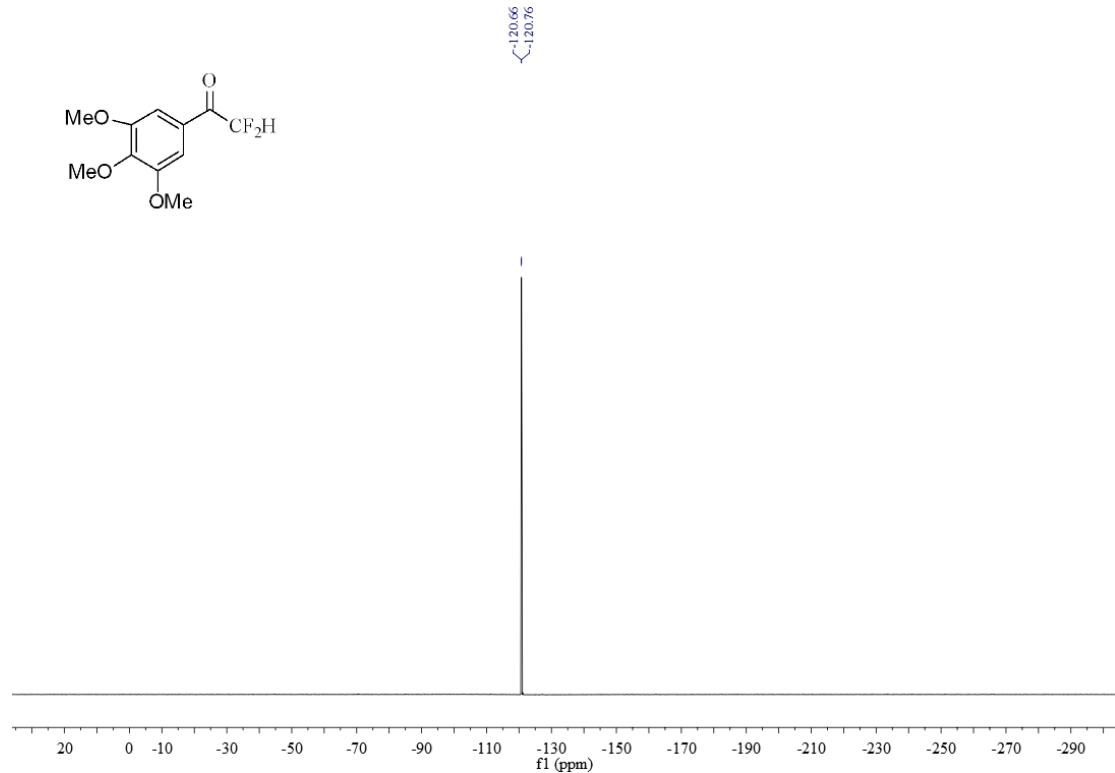
¹⁹F NMR of **4d**



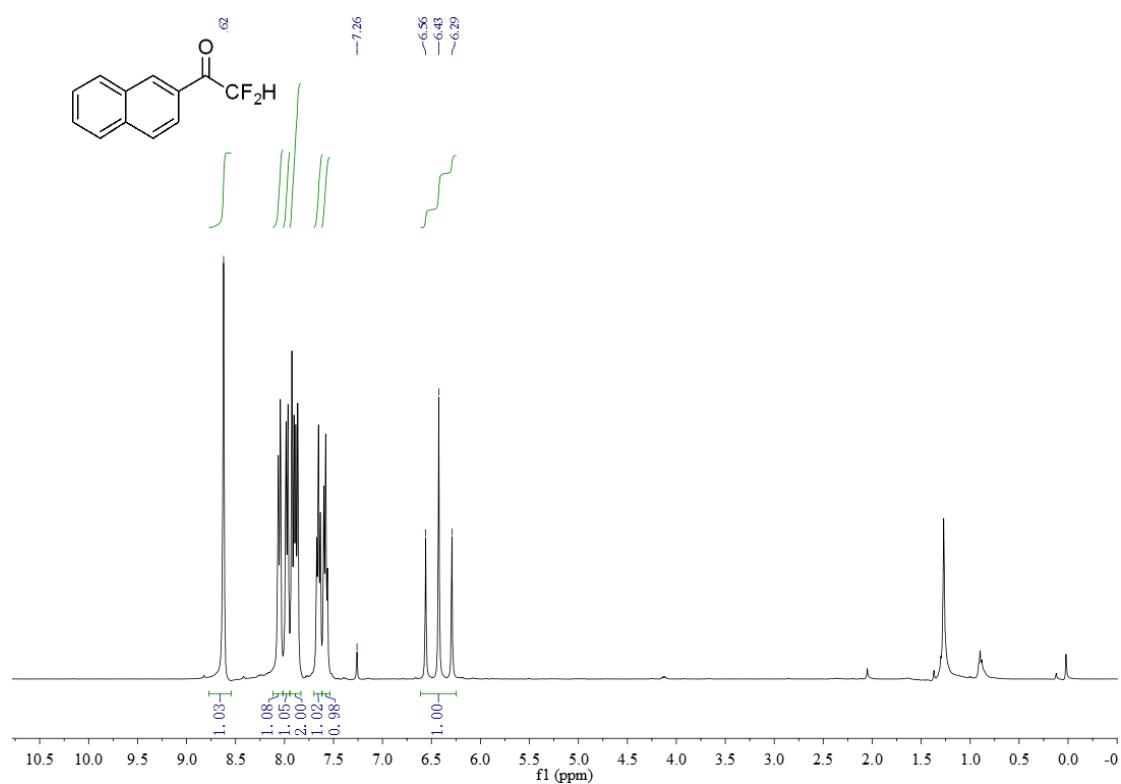
¹H NMR of **4e**



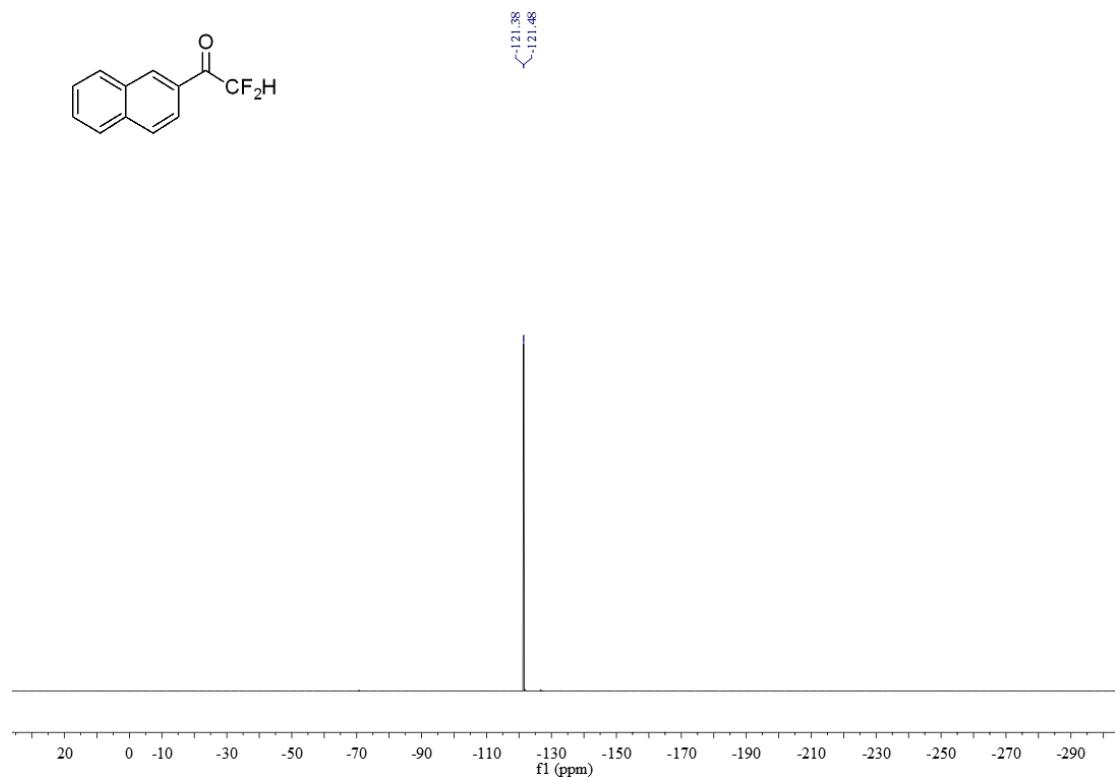
¹⁹F NMR of **4e**



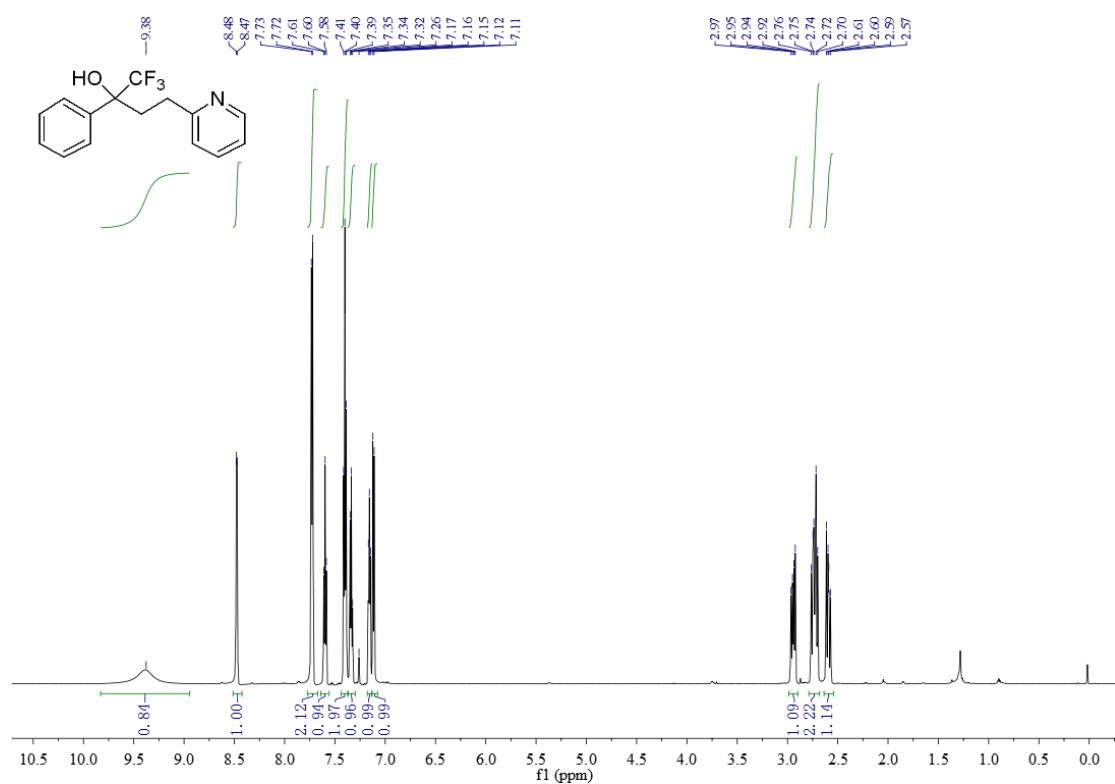
¹H NMR of **4f**



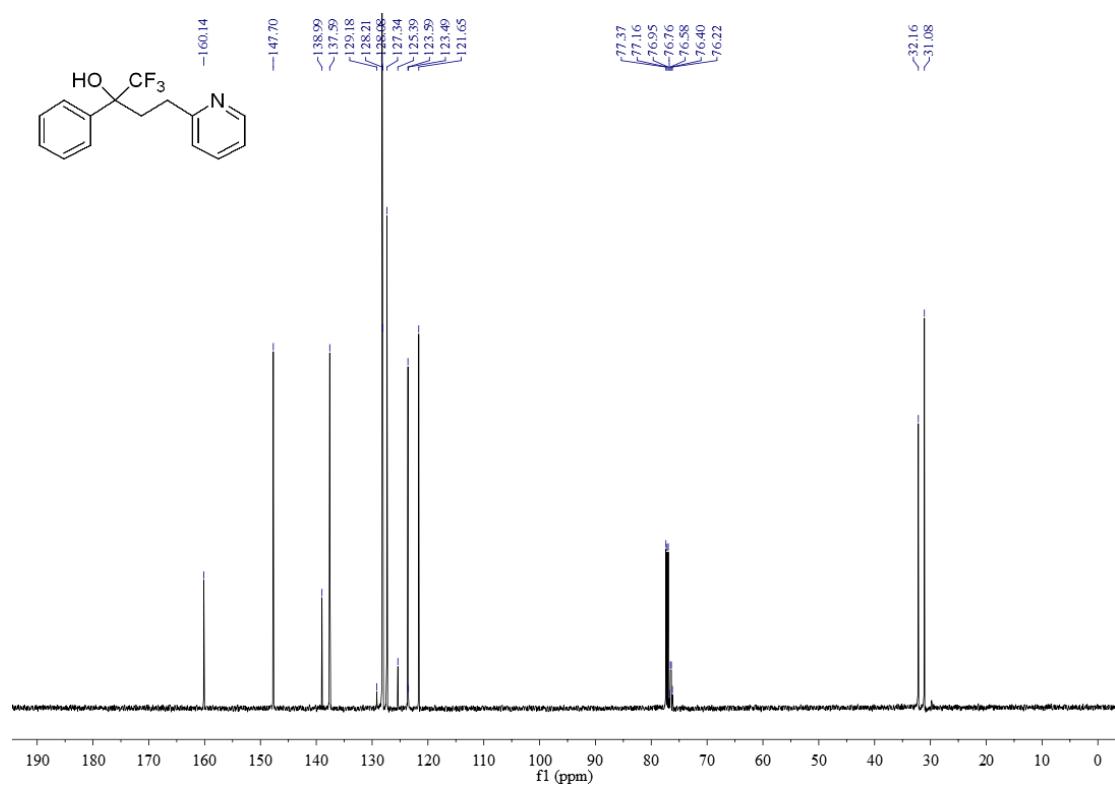
¹⁹F NMR of **4f**



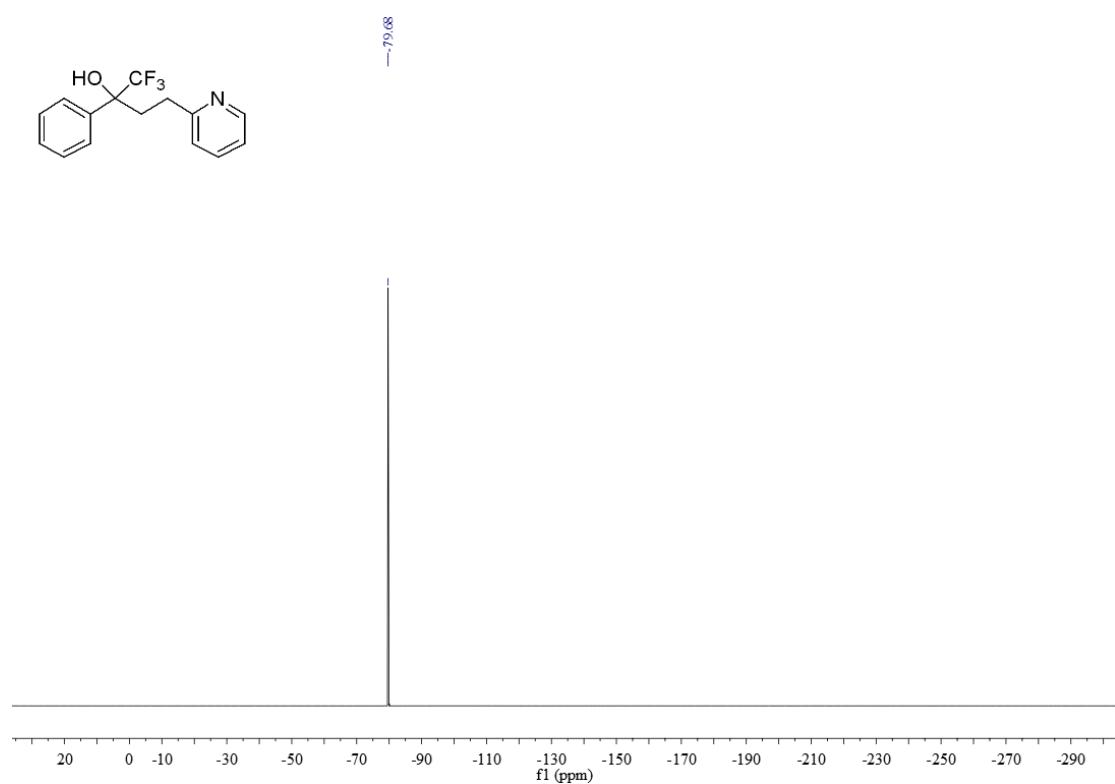
¹H NMR of **3a**



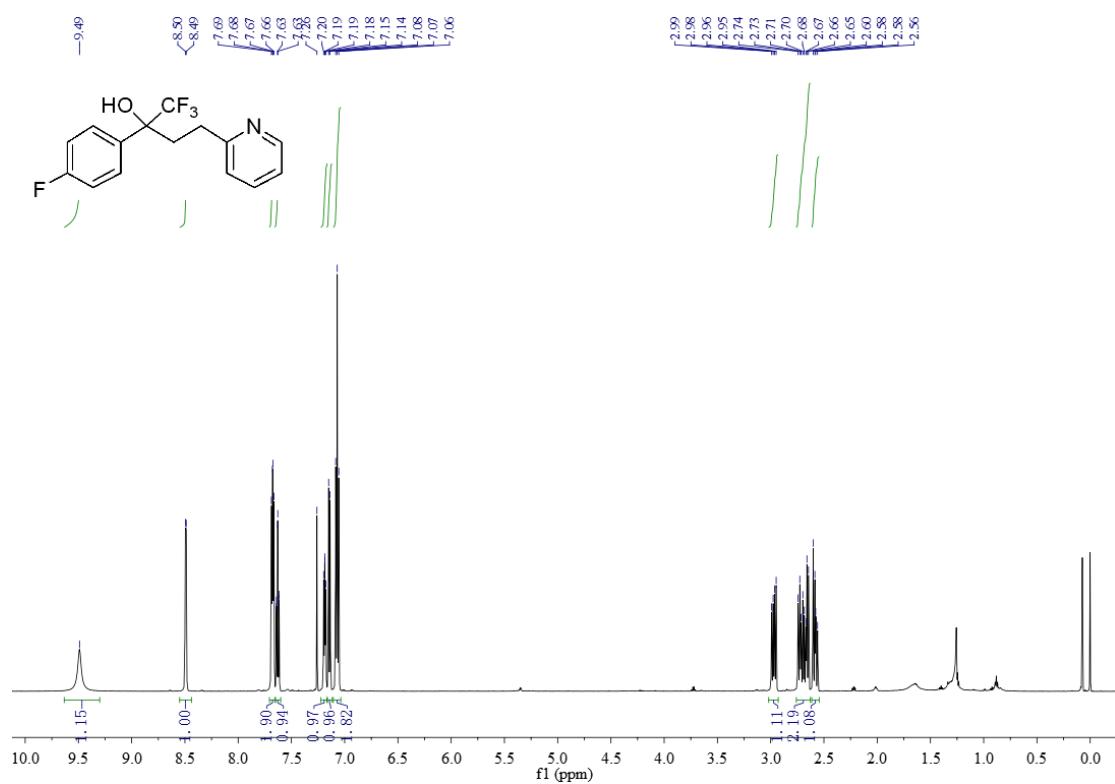
¹³C NMR of **3a**



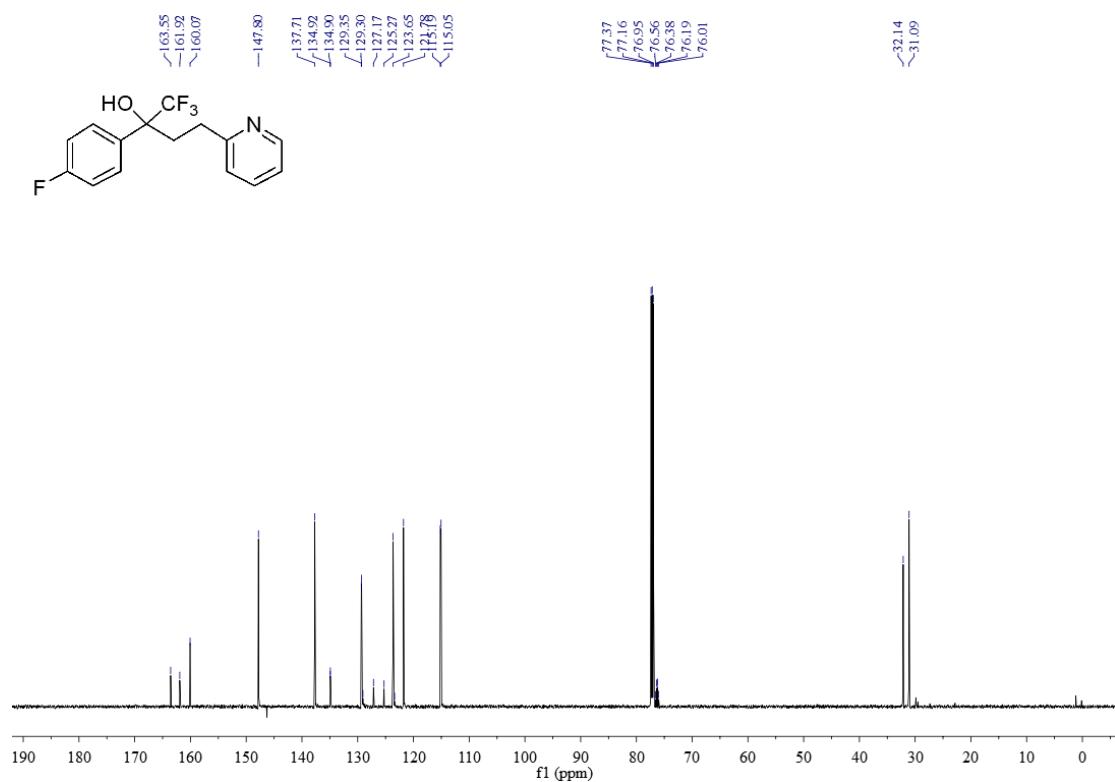
¹⁹F NMR of **3a**



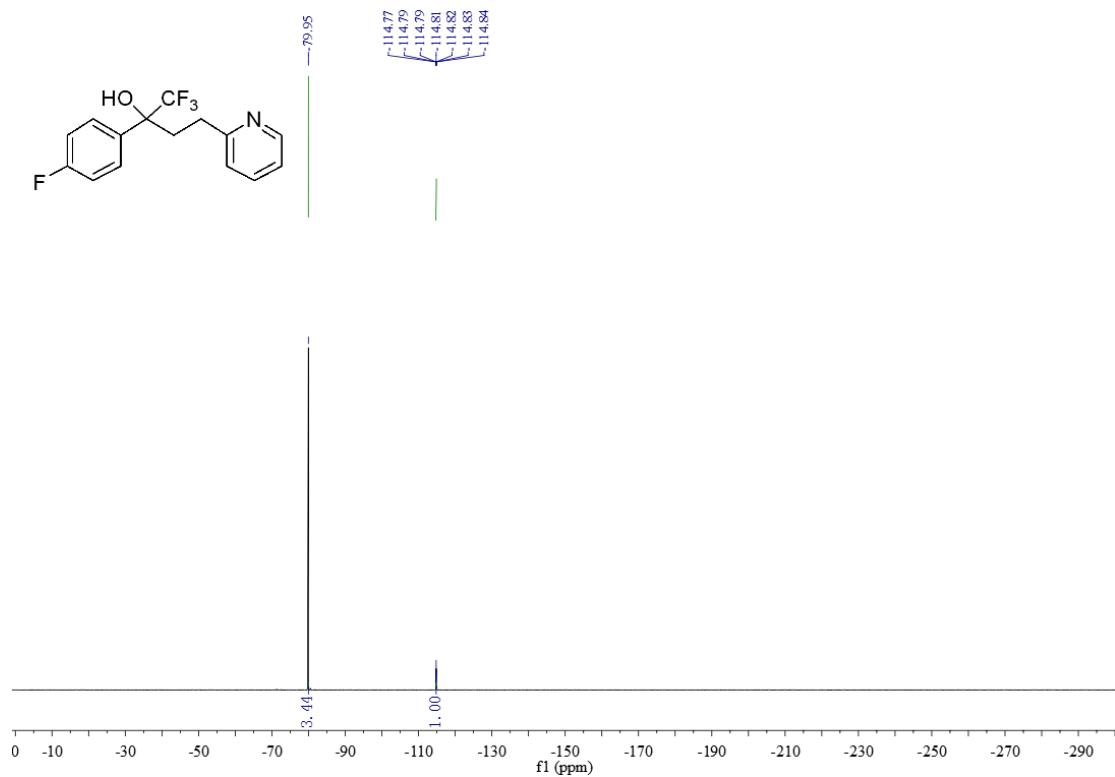
¹H NMR of **3b**



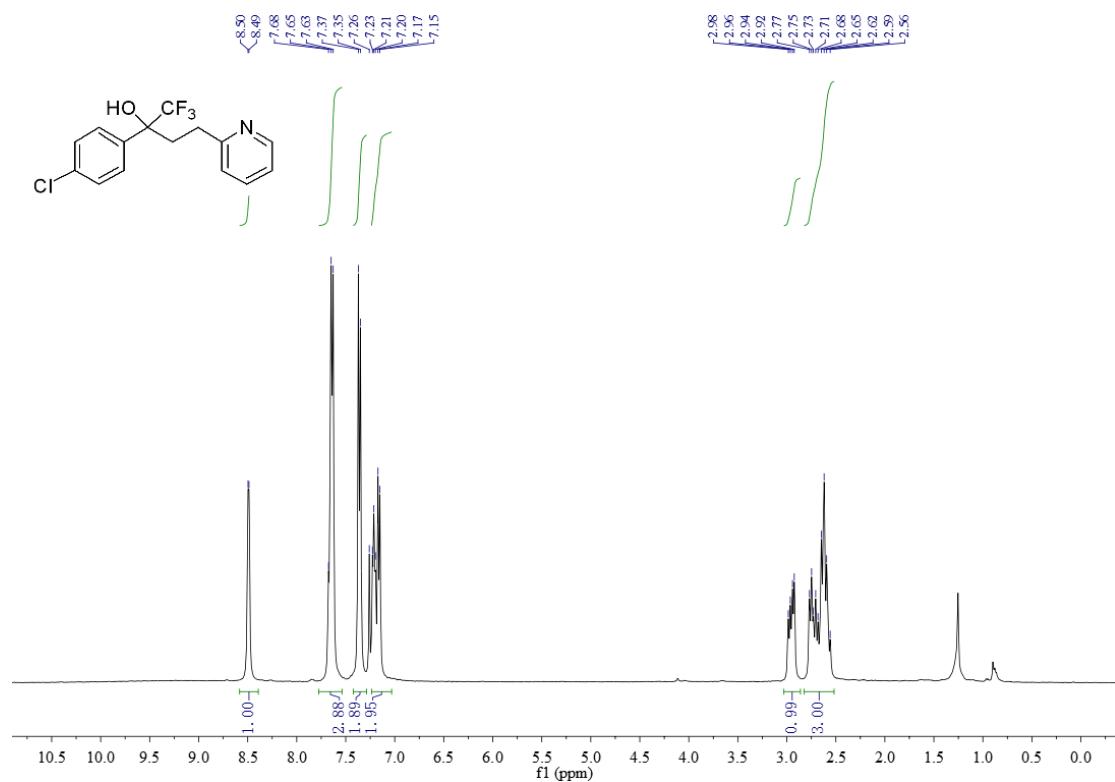
¹³C NMR of **3b**



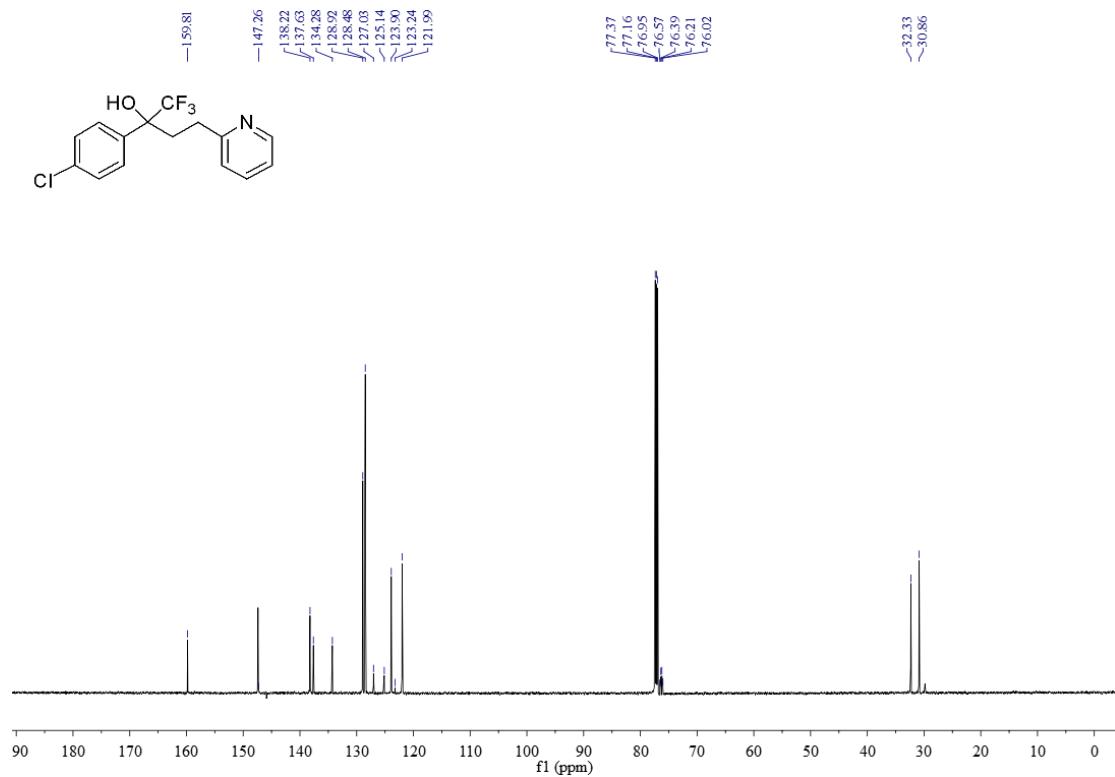
¹⁹F NMR of **3b**



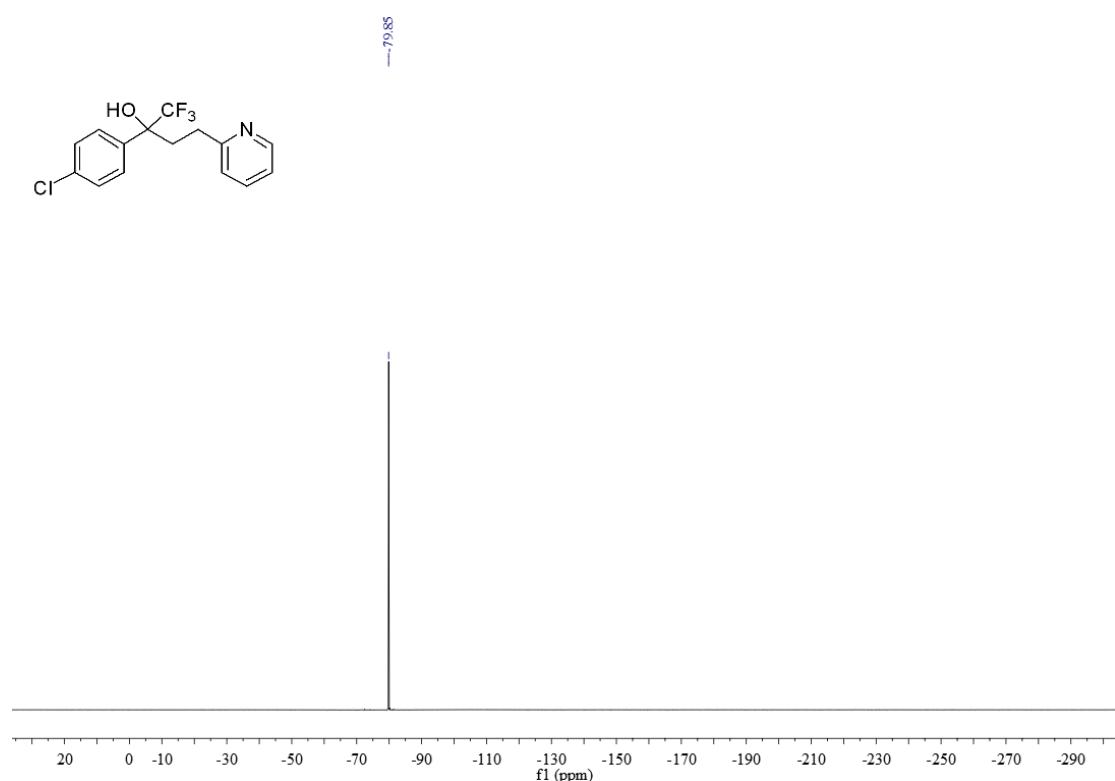
¹H NMR of **3c**



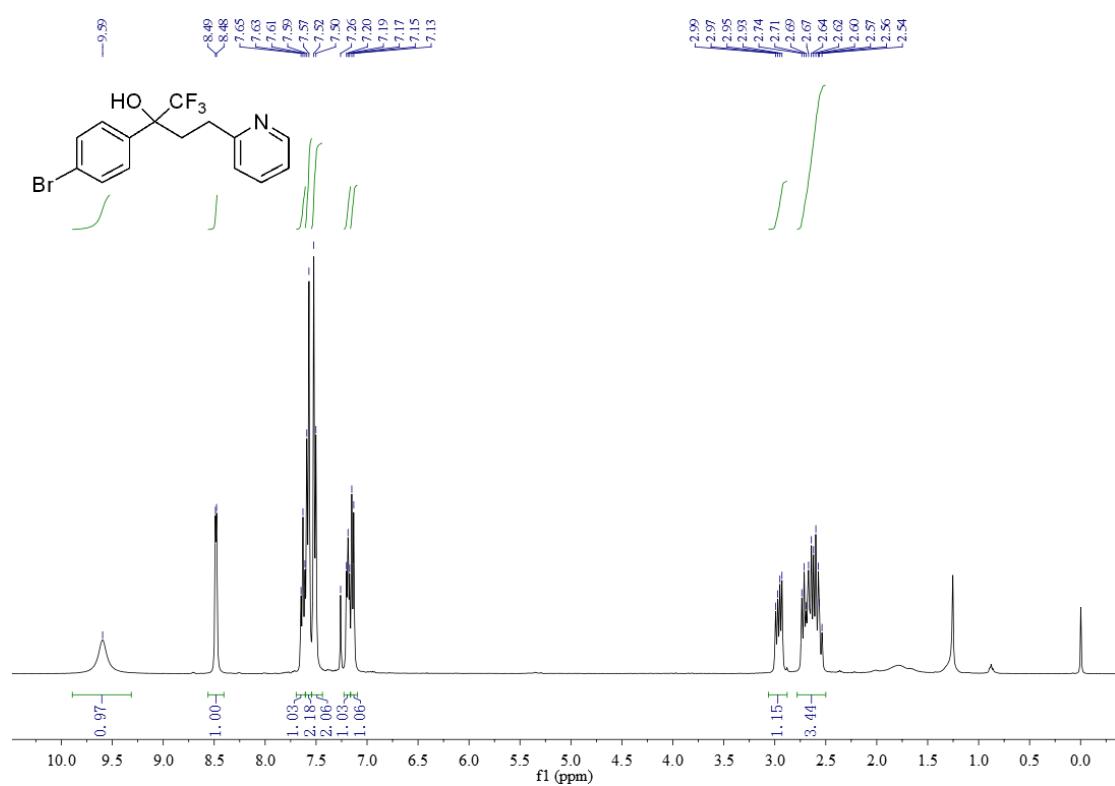
¹³C NMR of **3c**



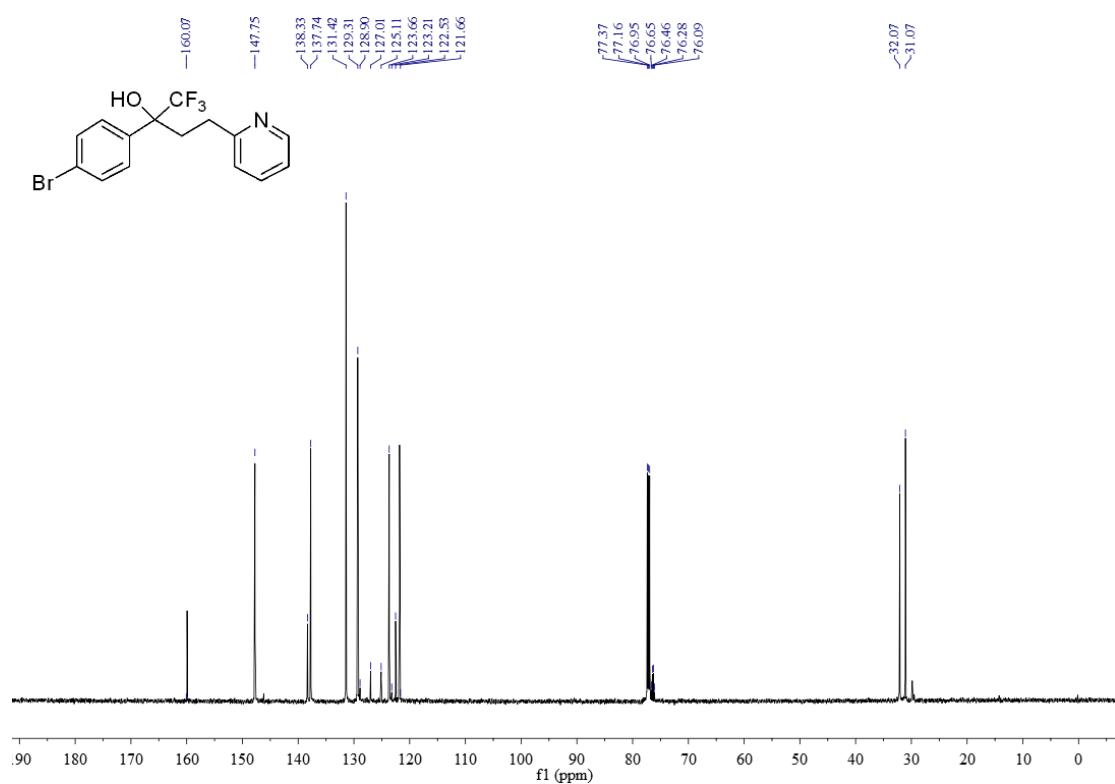
¹⁹F NMR of **3c**



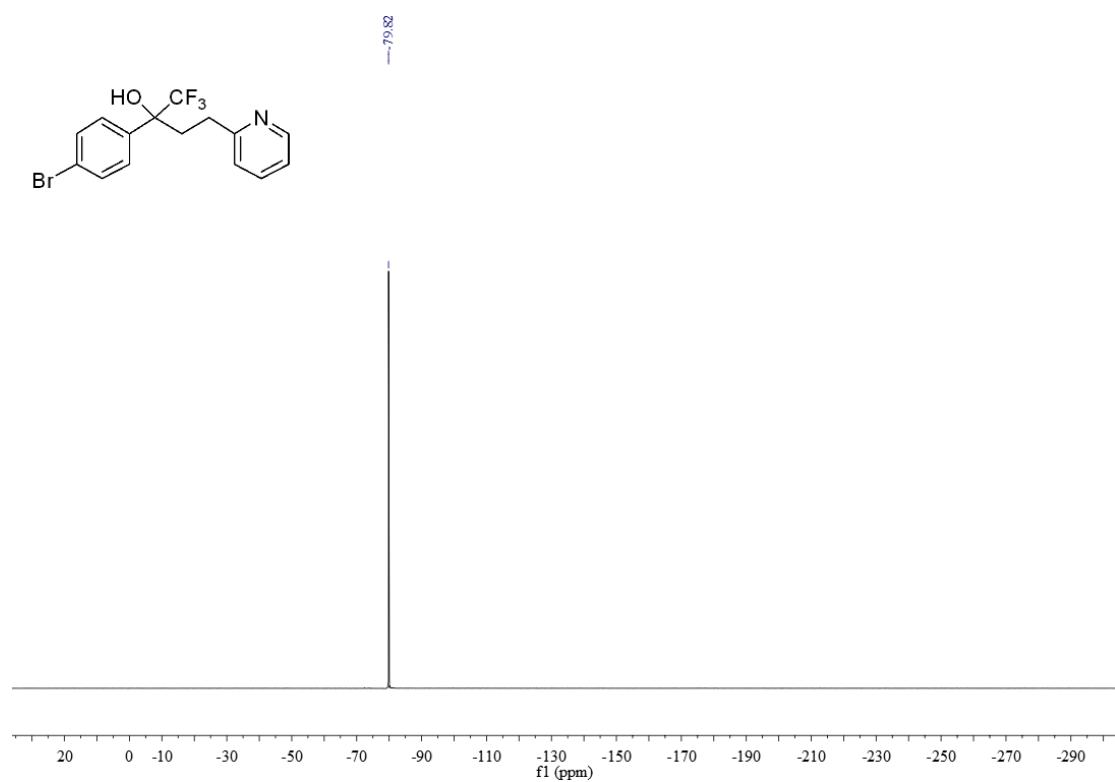
¹H NMR of **3d**



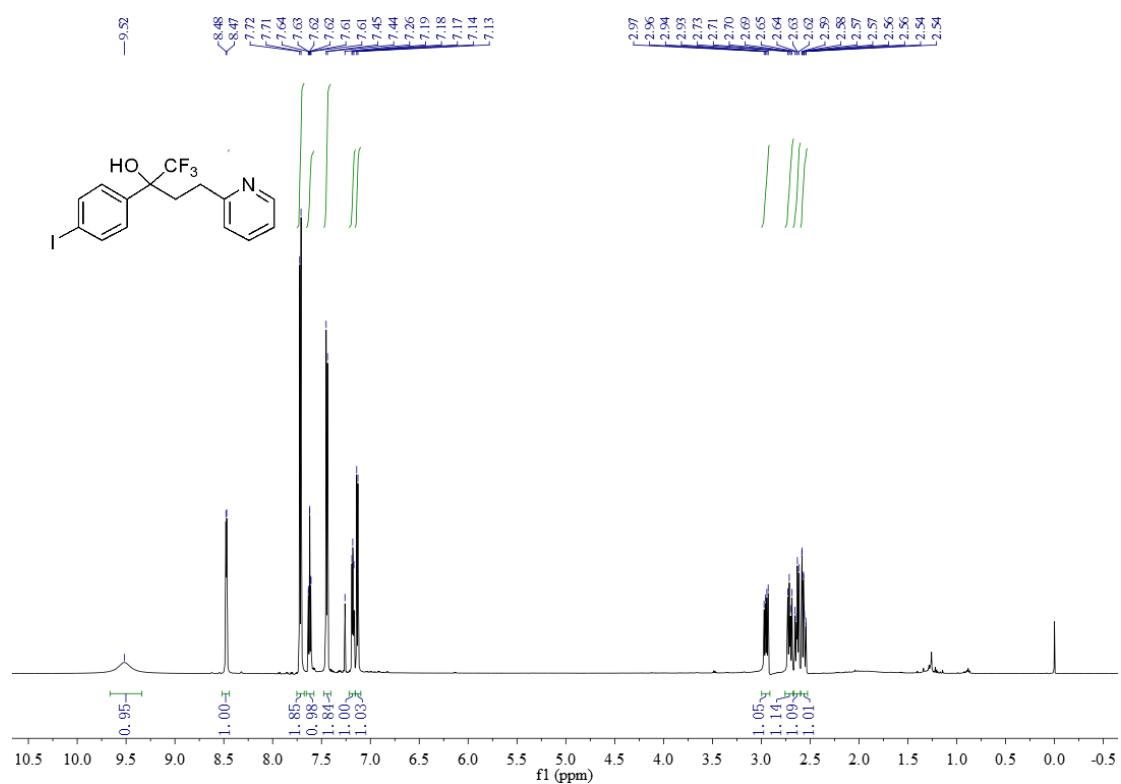
¹³C NMR of **3d**



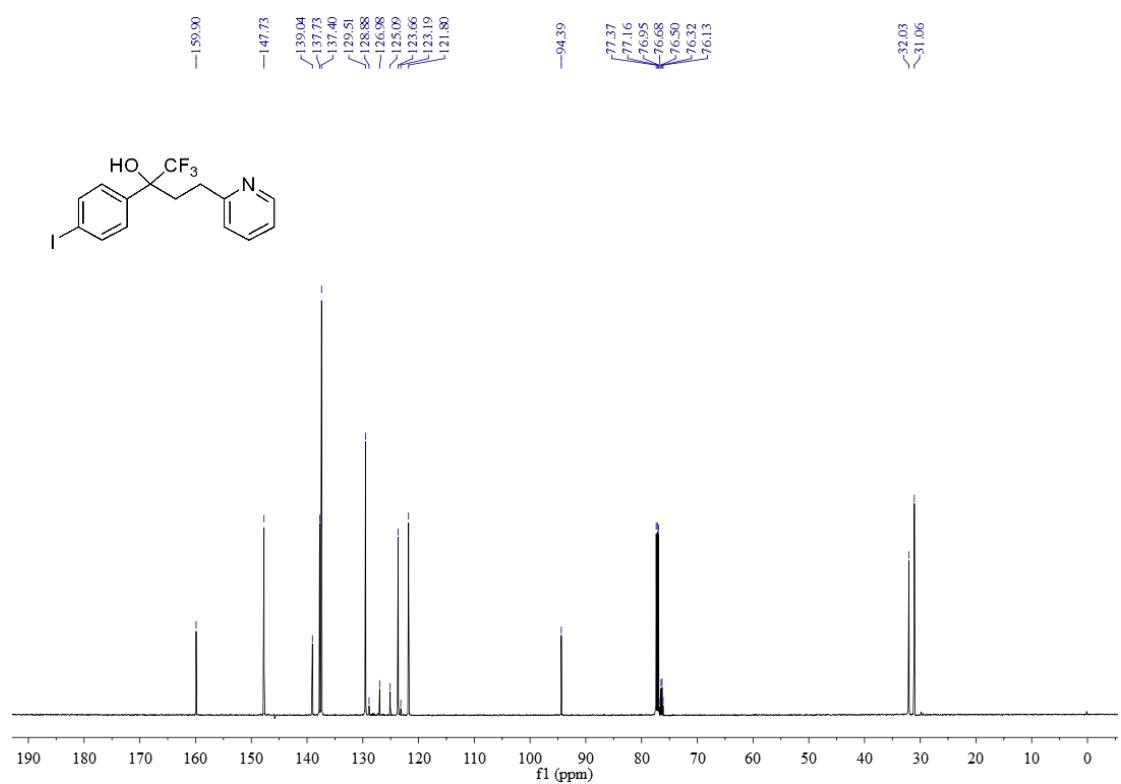
¹⁹F NMR of **3d**



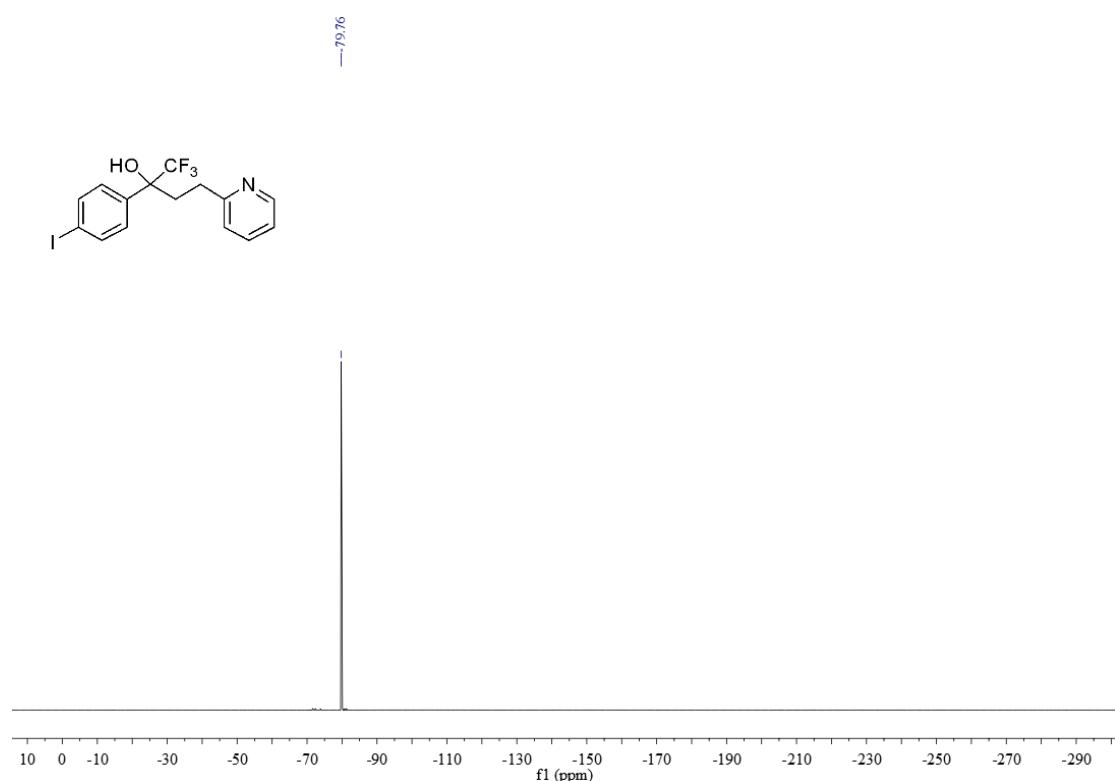
¹H NMR of **3e**



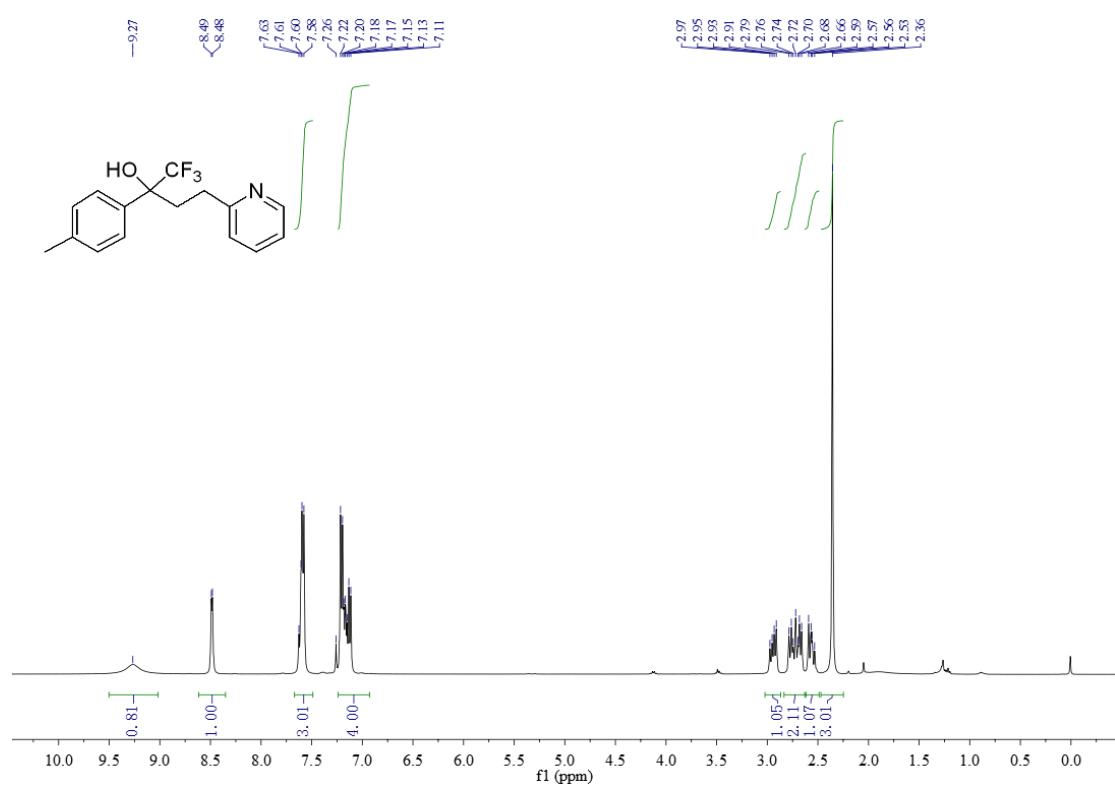
¹³C NMR of **3e**



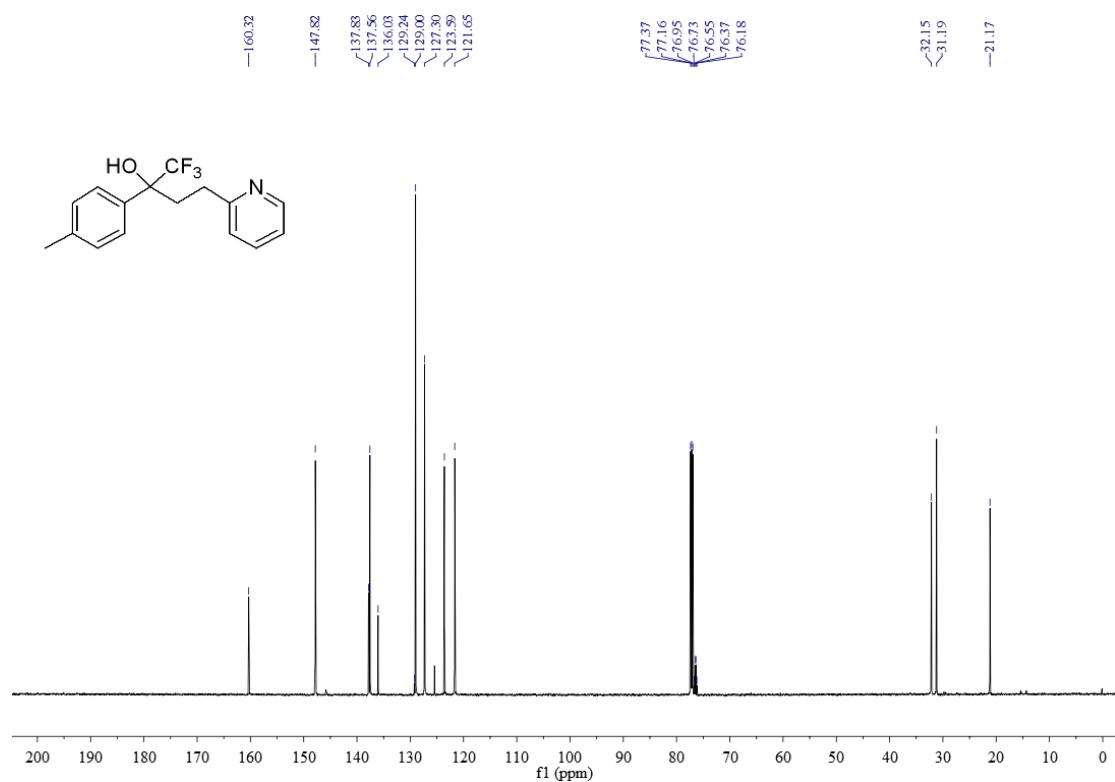
¹⁹F NMR of **3e**



¹H NMR of **3f**



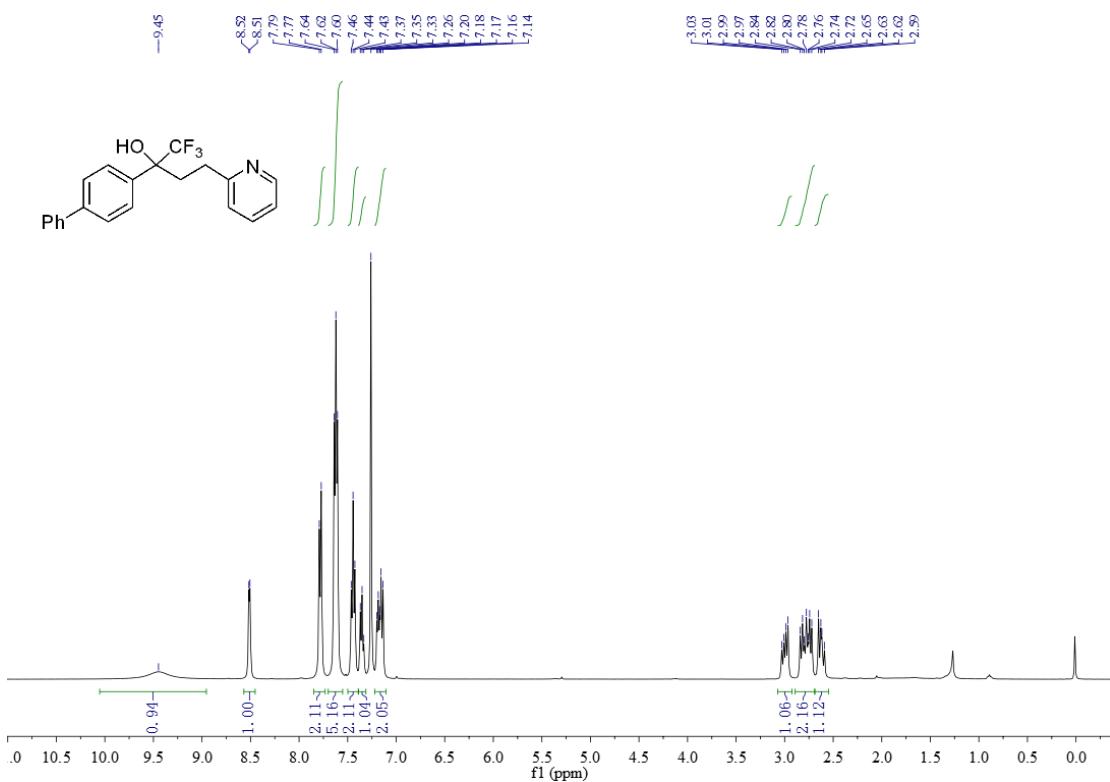
¹³C NMR of **3f**



¹⁹F NMR of **3f**



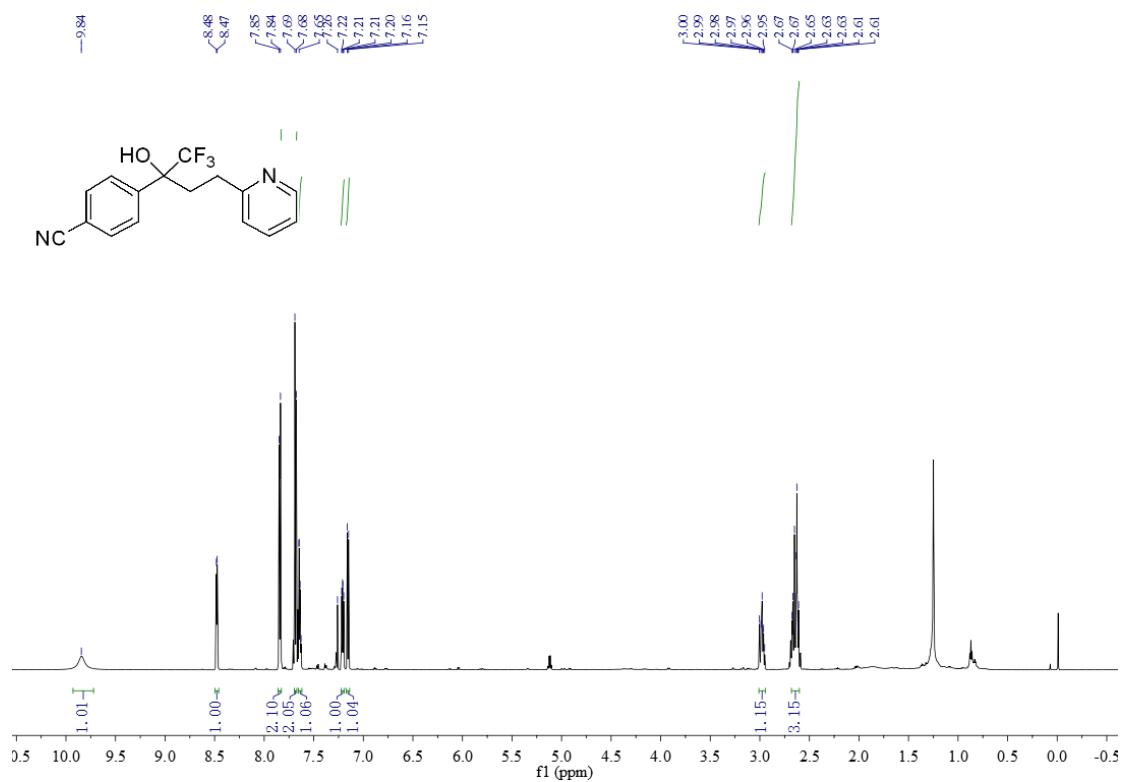
¹H NMR of **3g**



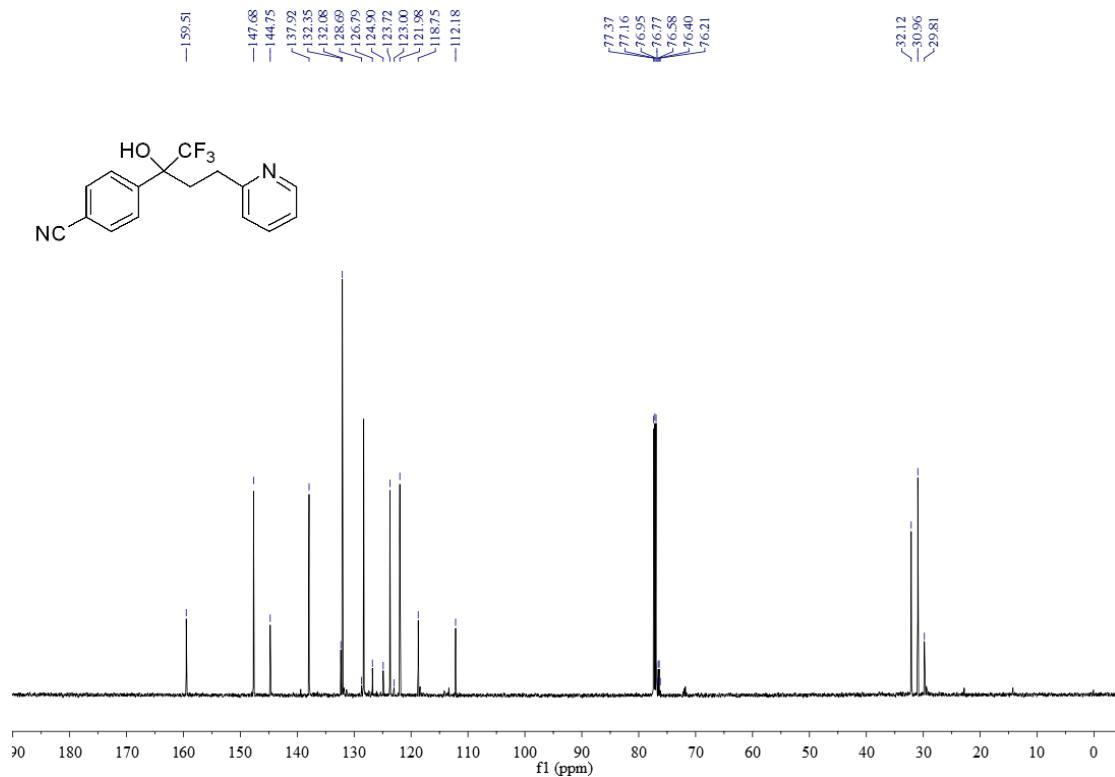
¹⁹F NMR of **3g**



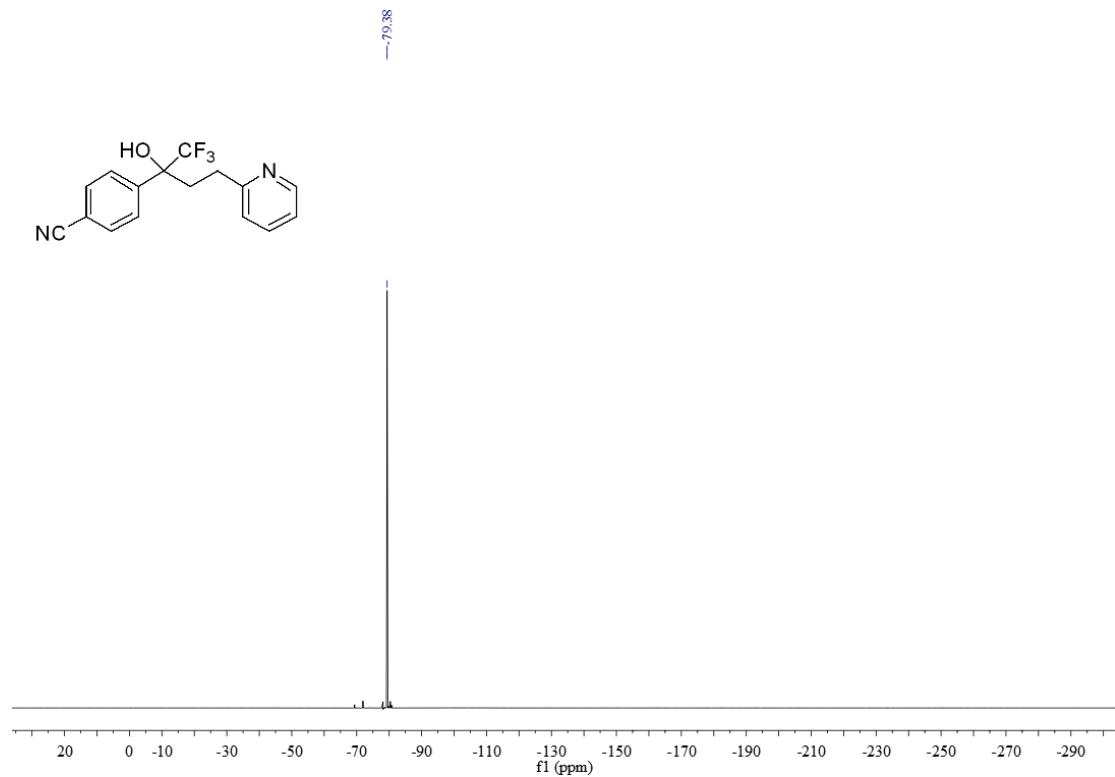
¹H NMR of **3h**



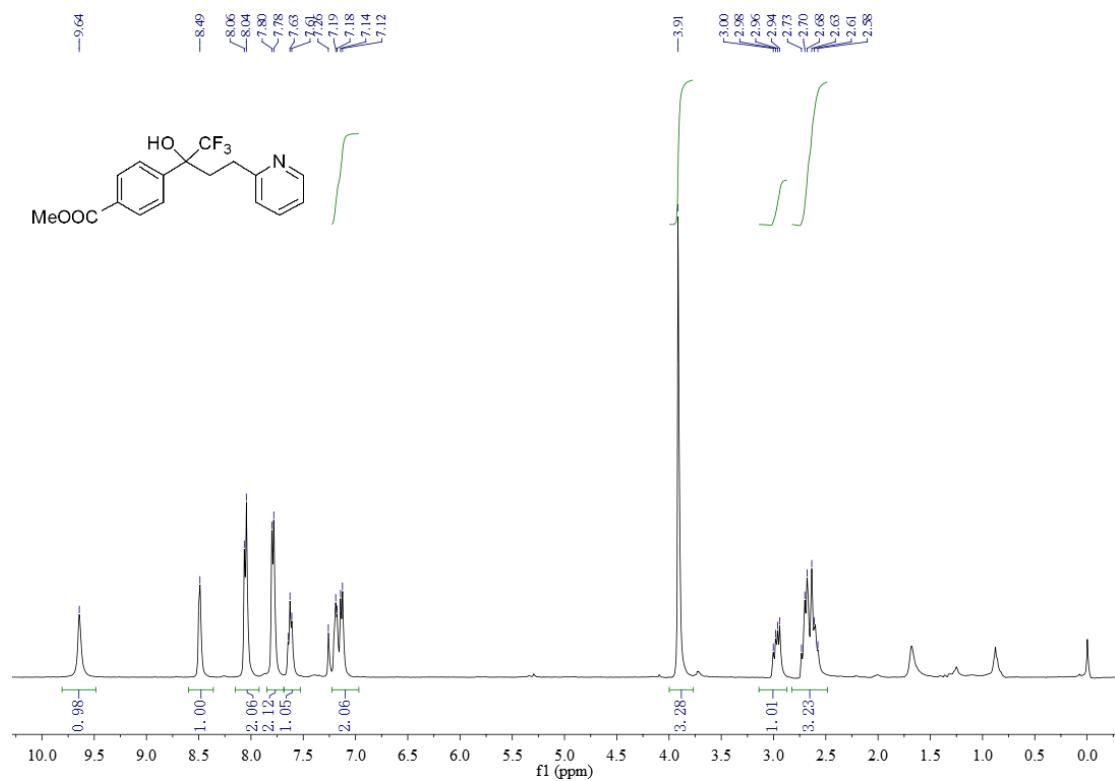
¹³C NMR of **3h**



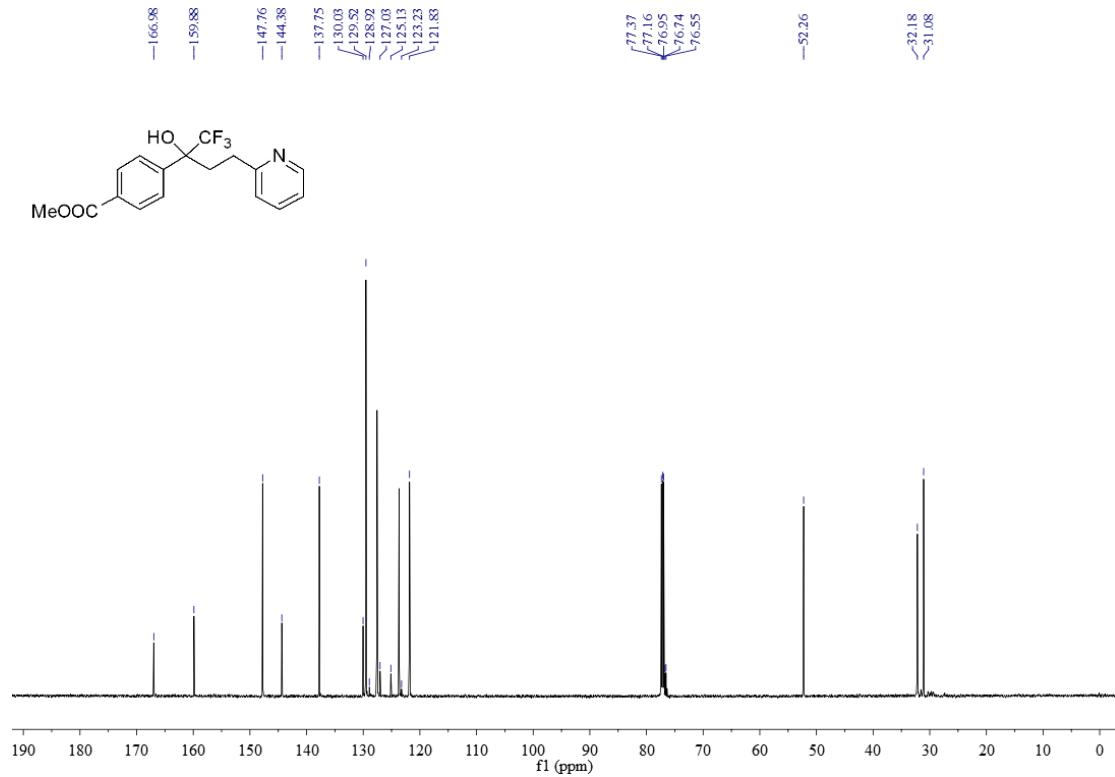
¹⁹F NMR of **3h**



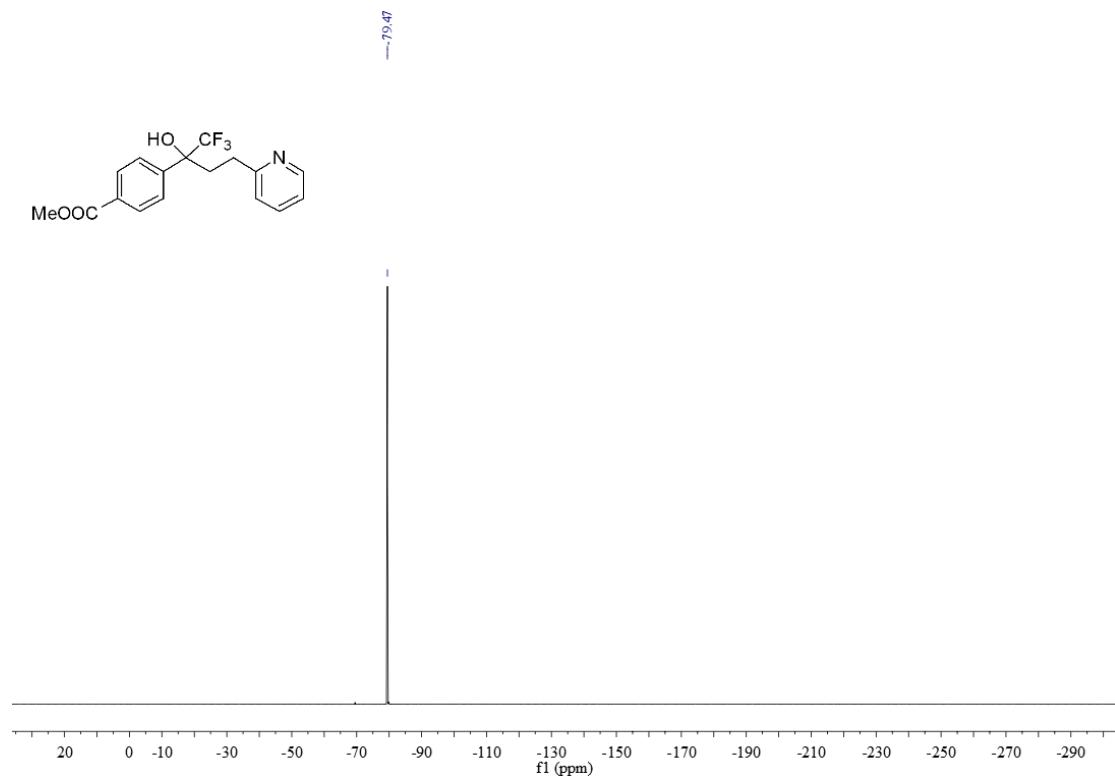
¹H NMR of **3i**



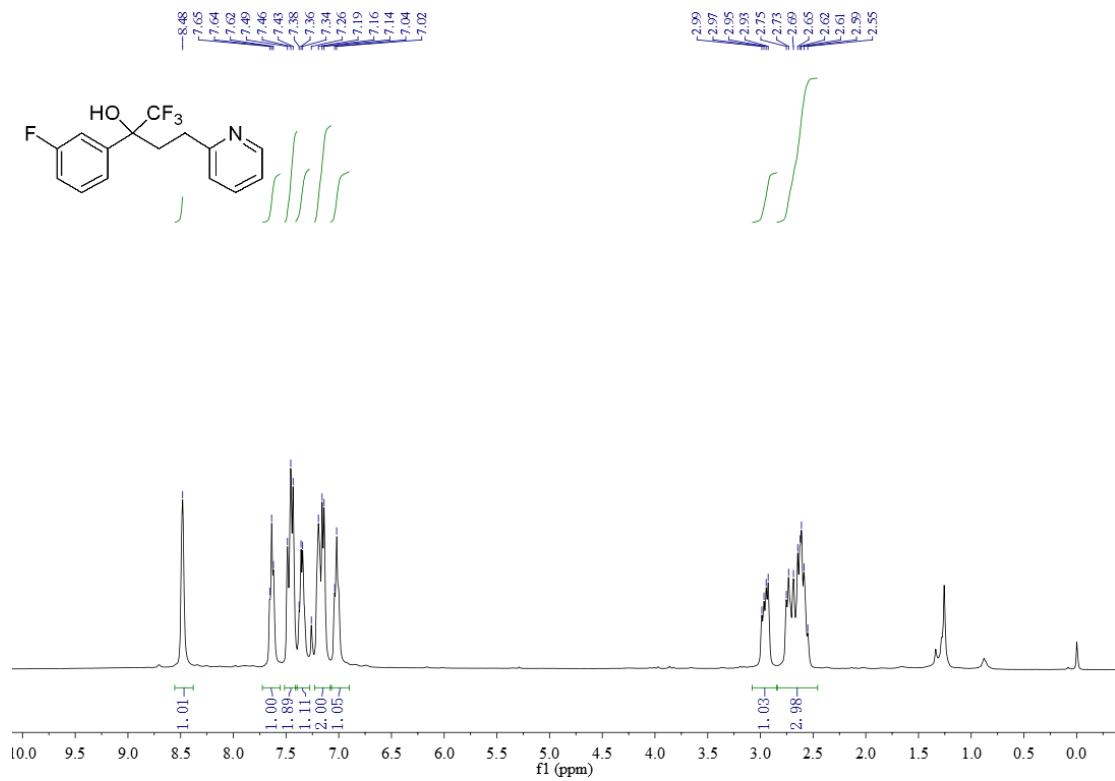
¹³C NMR of **3i**



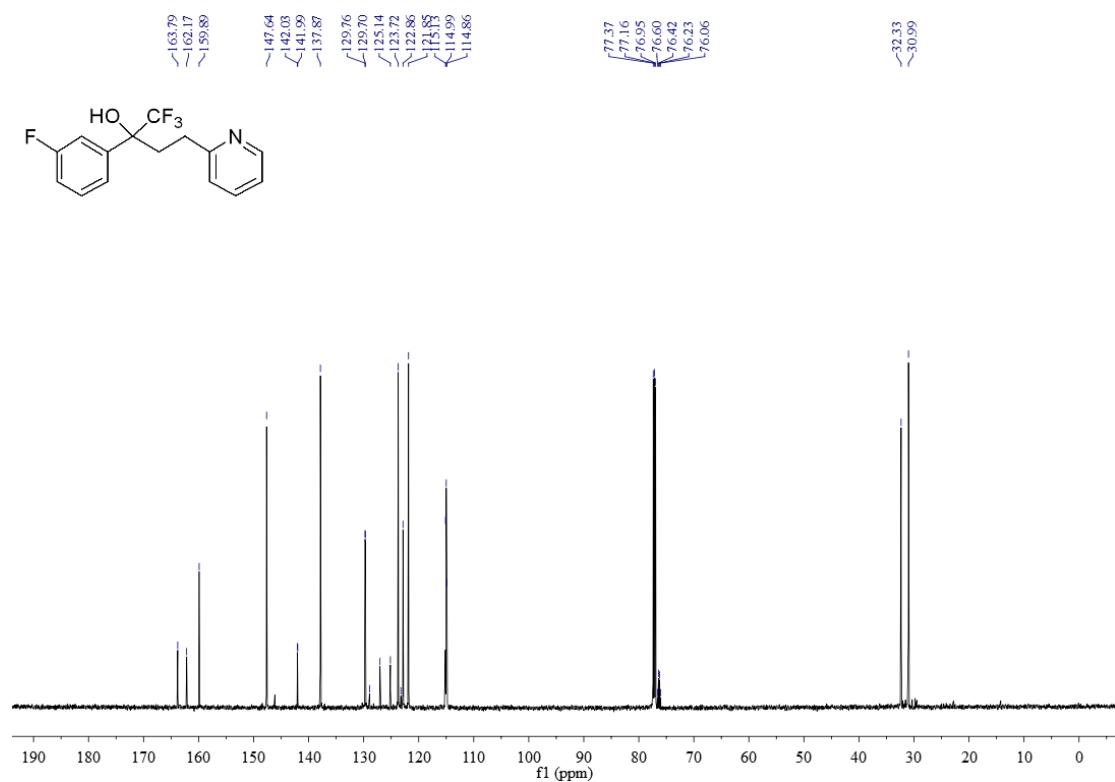
¹⁹F NMR of **3i**



¹H NMR of **3j**



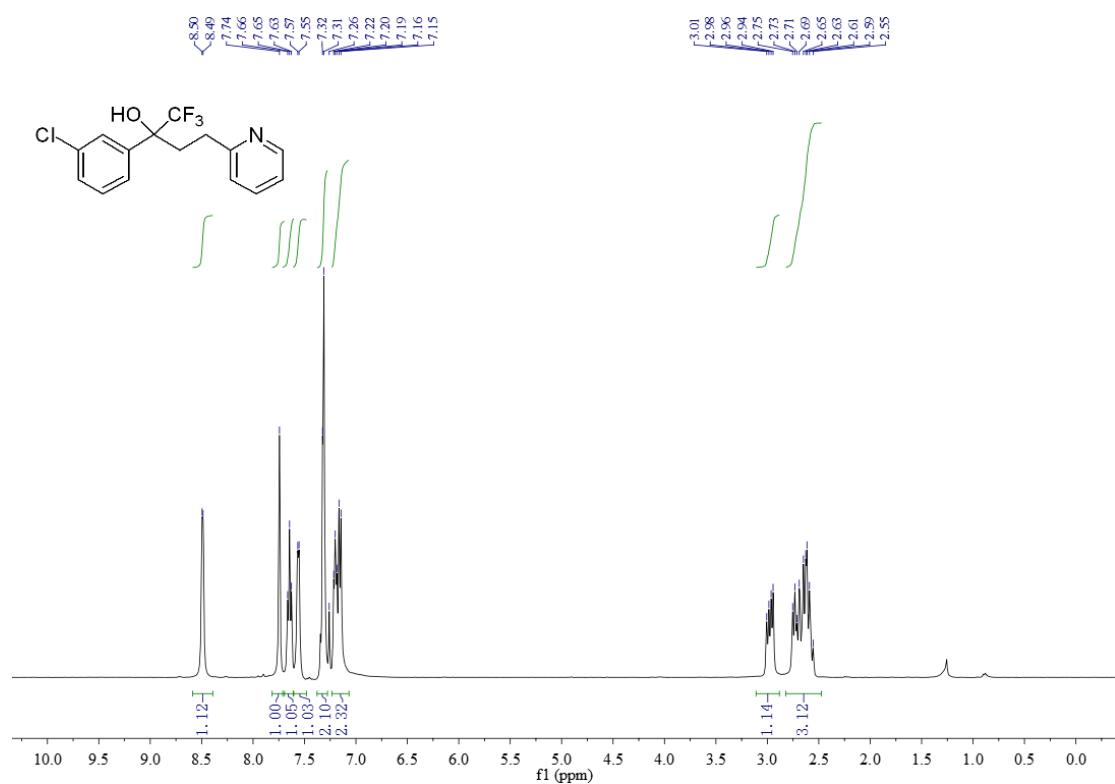
¹³C NMR of **3j**



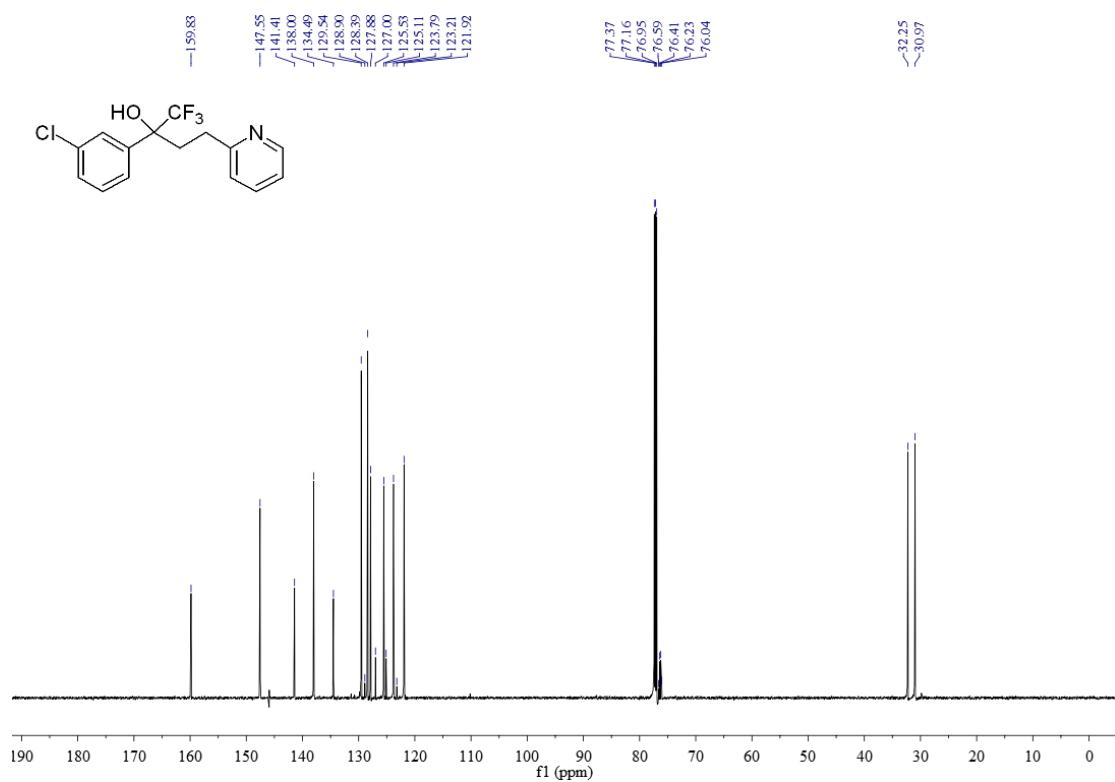
¹⁹F NMR of **3j**



¹H NMR of **3k**



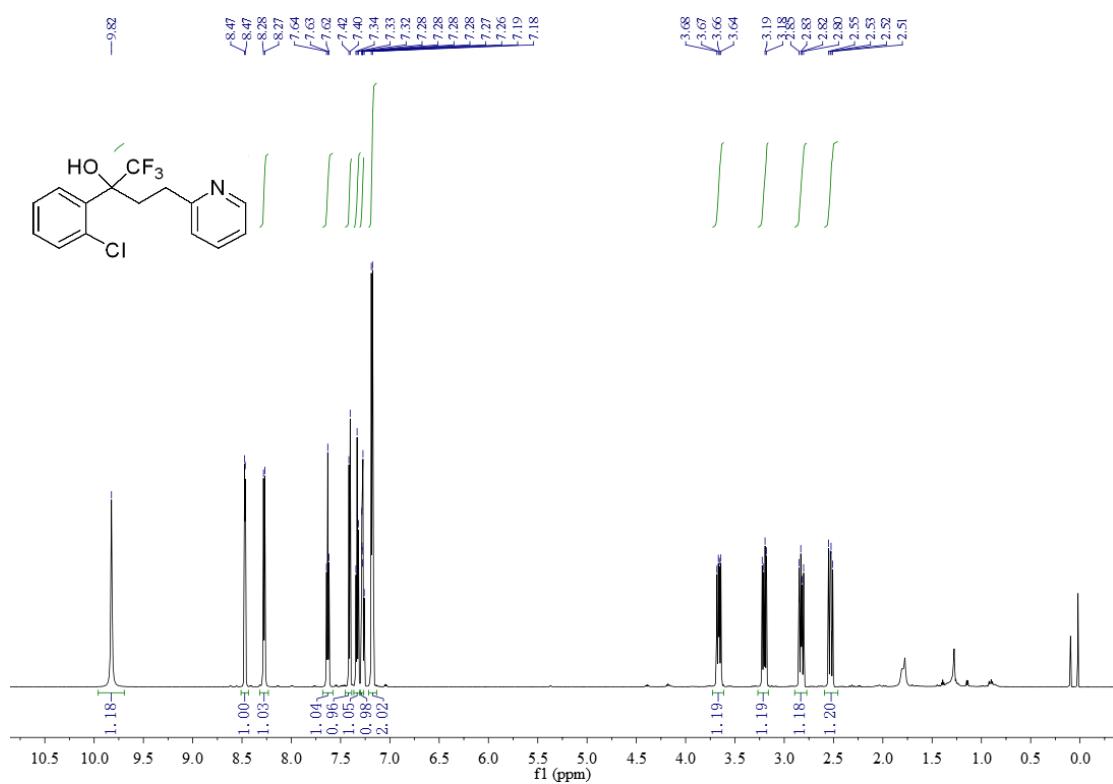
¹³C NMR of **3k**



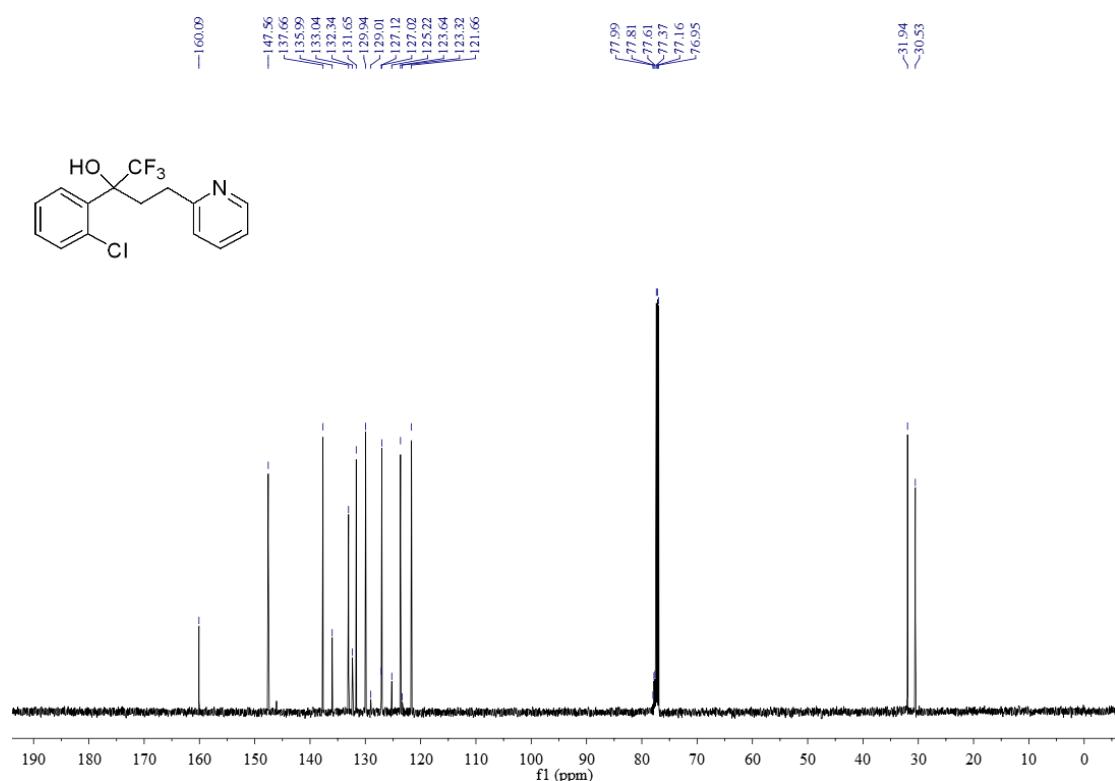
¹⁹F NMR of **3k**



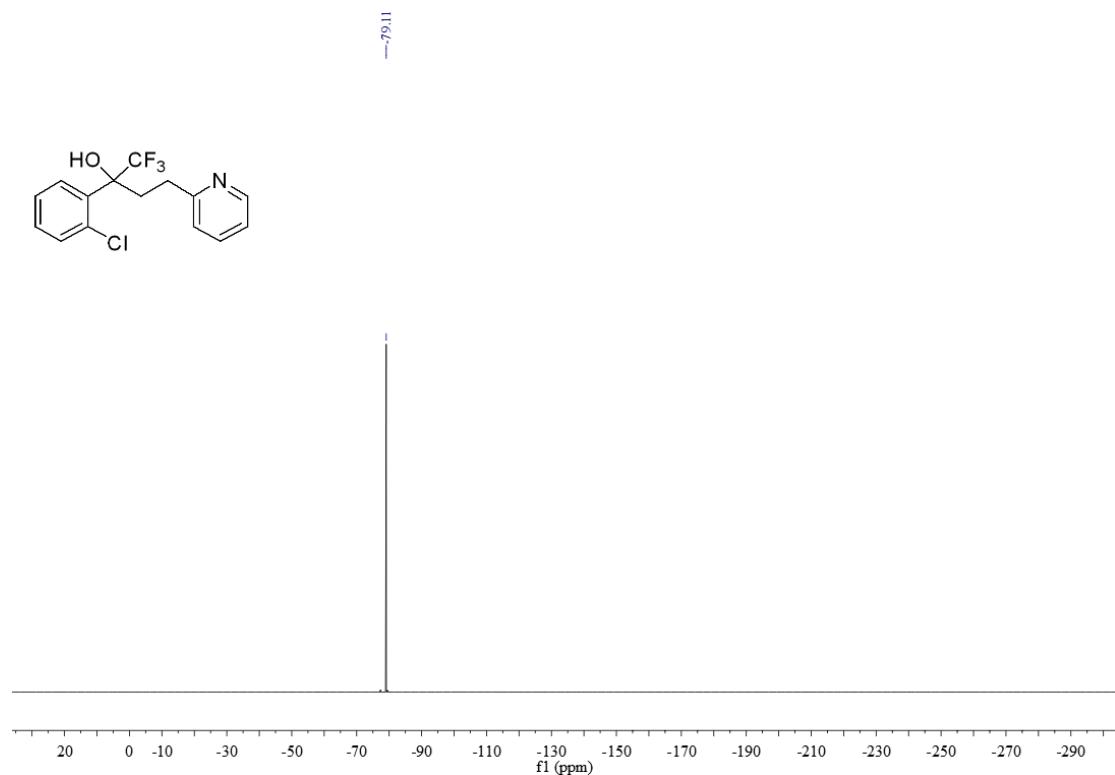
¹H NMR of **3l**



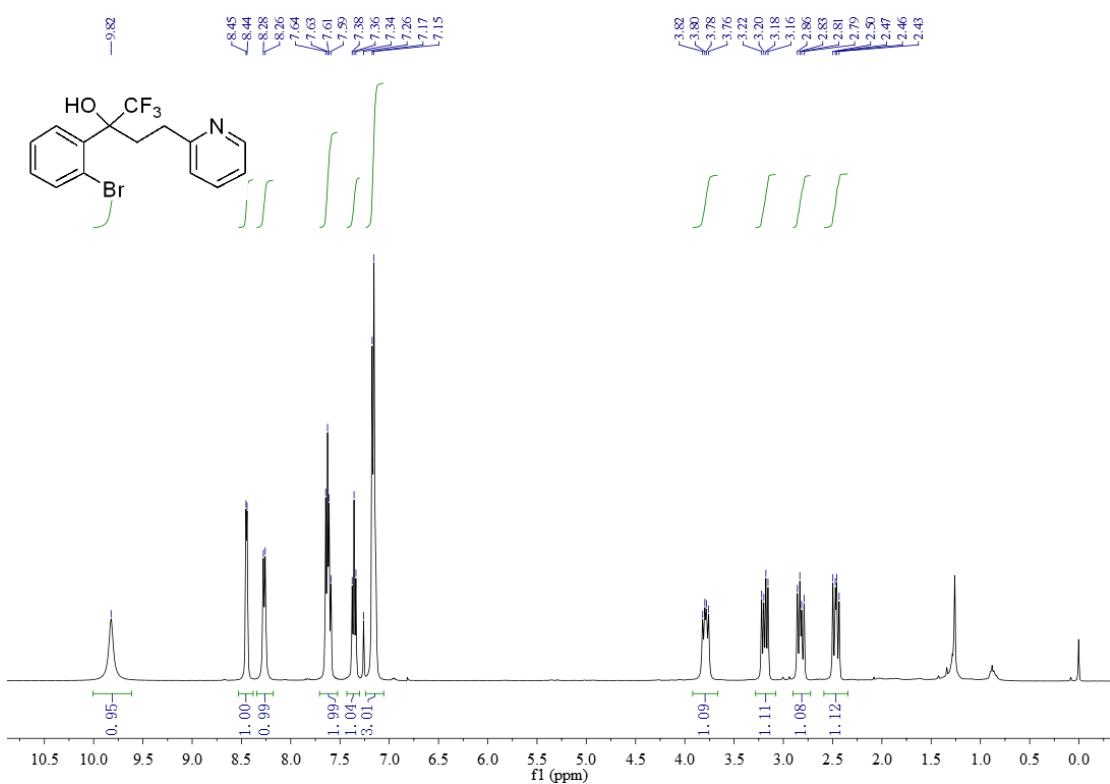
¹³C NMR of **3l**



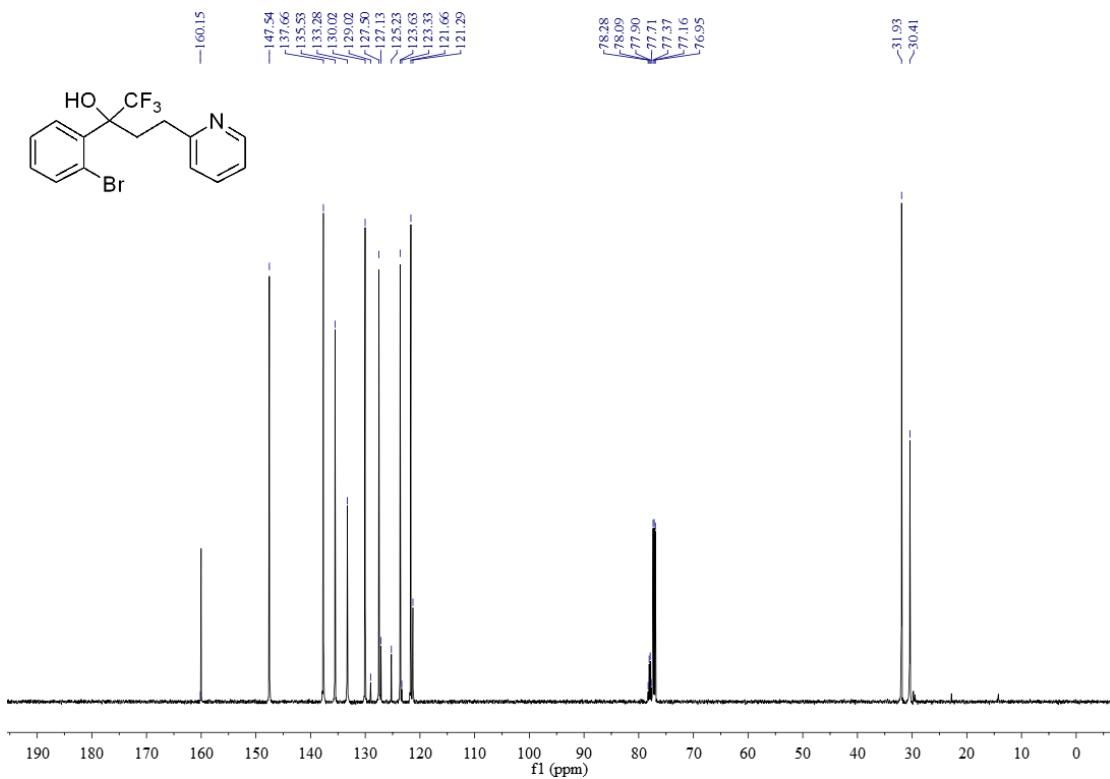
¹⁹F NMR of **3l**



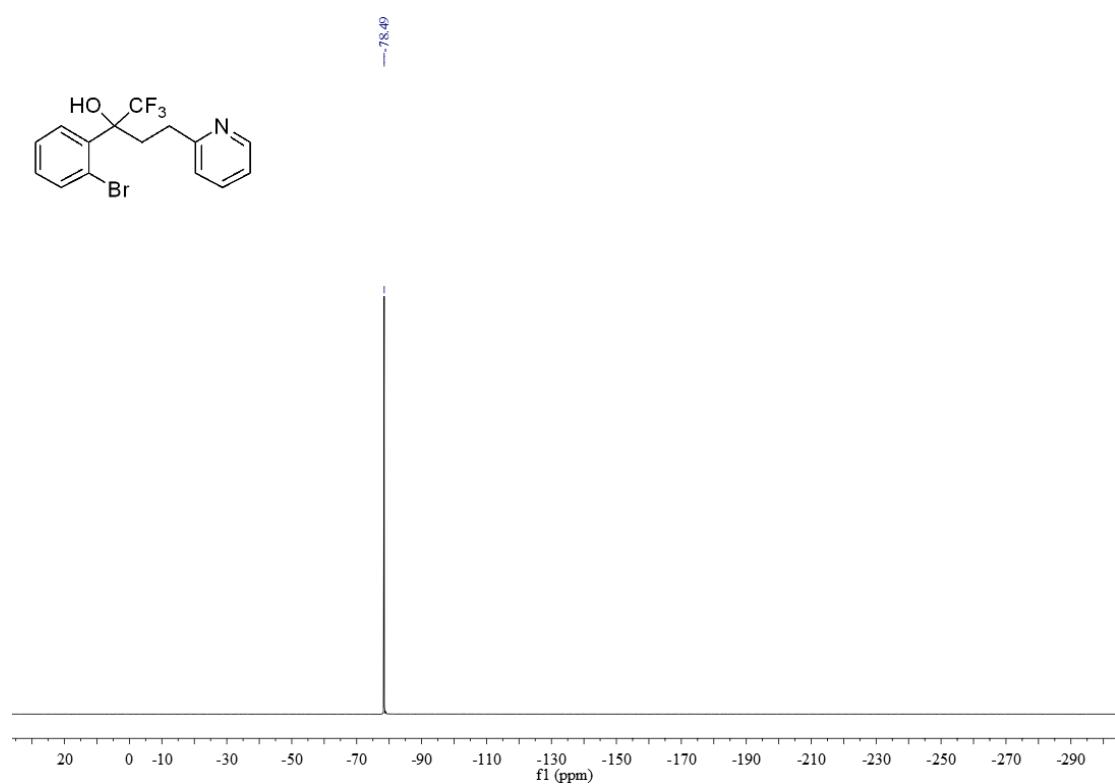
¹H NMR of **3m**



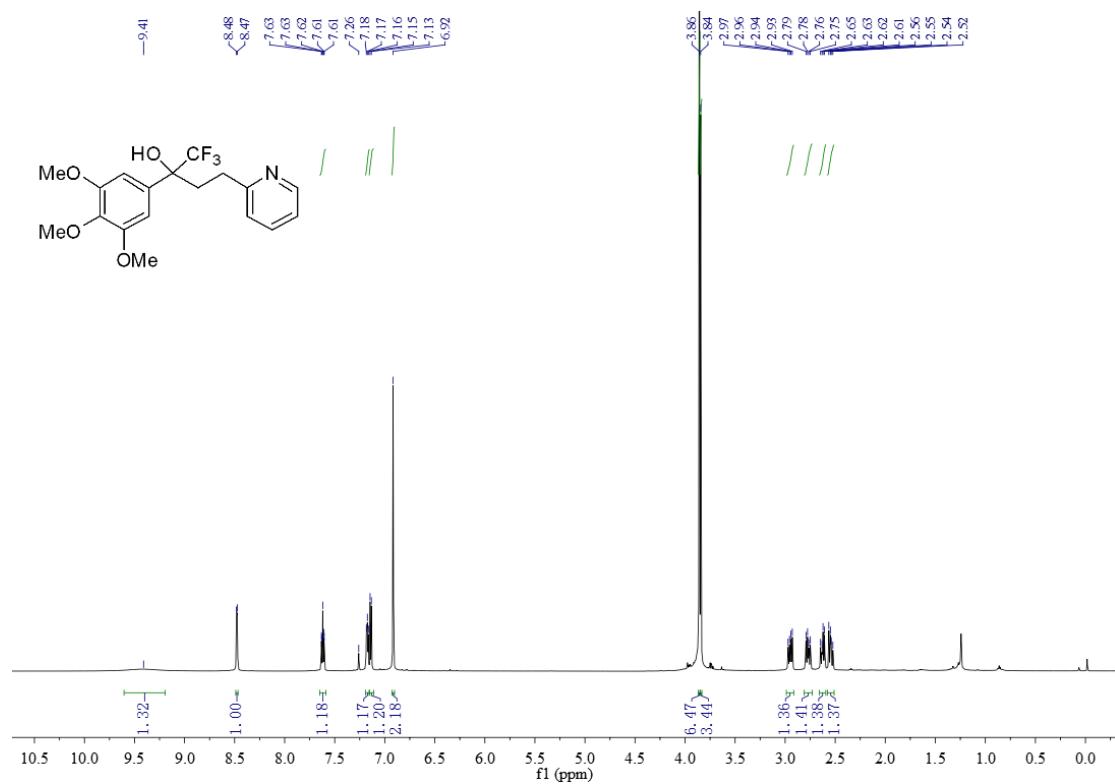
¹³C NMR of **3m**



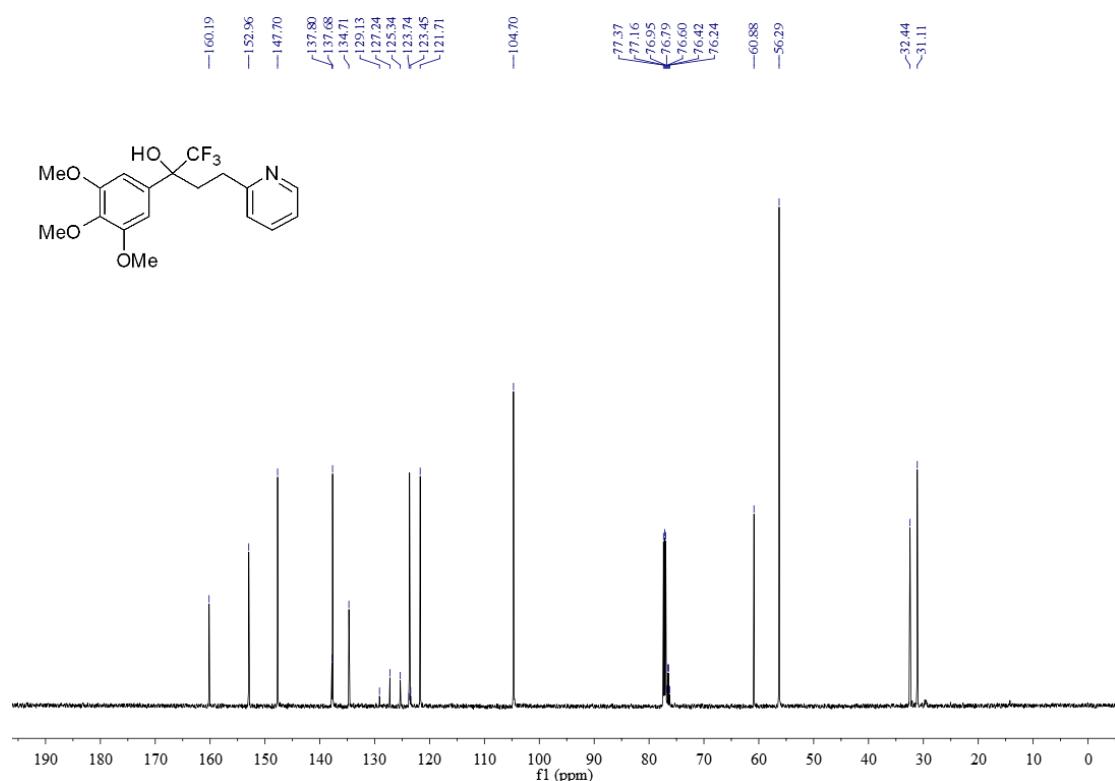
¹⁹F NMR of **3m**



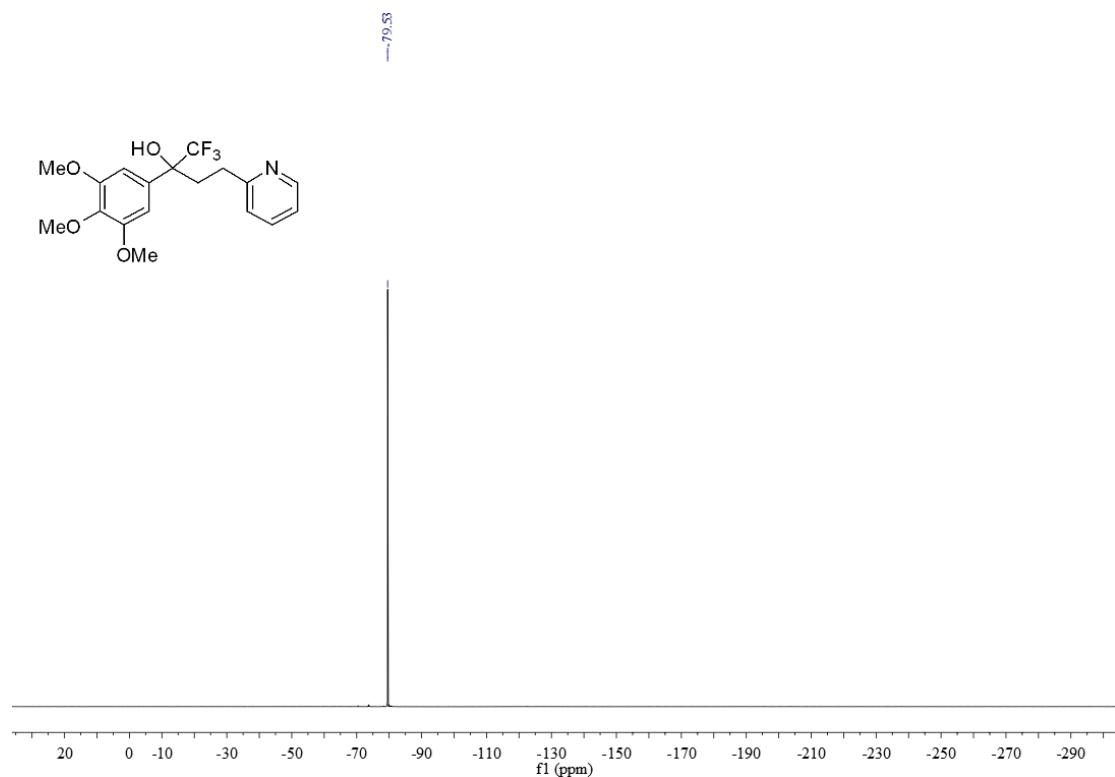
¹H NMR of **3n**



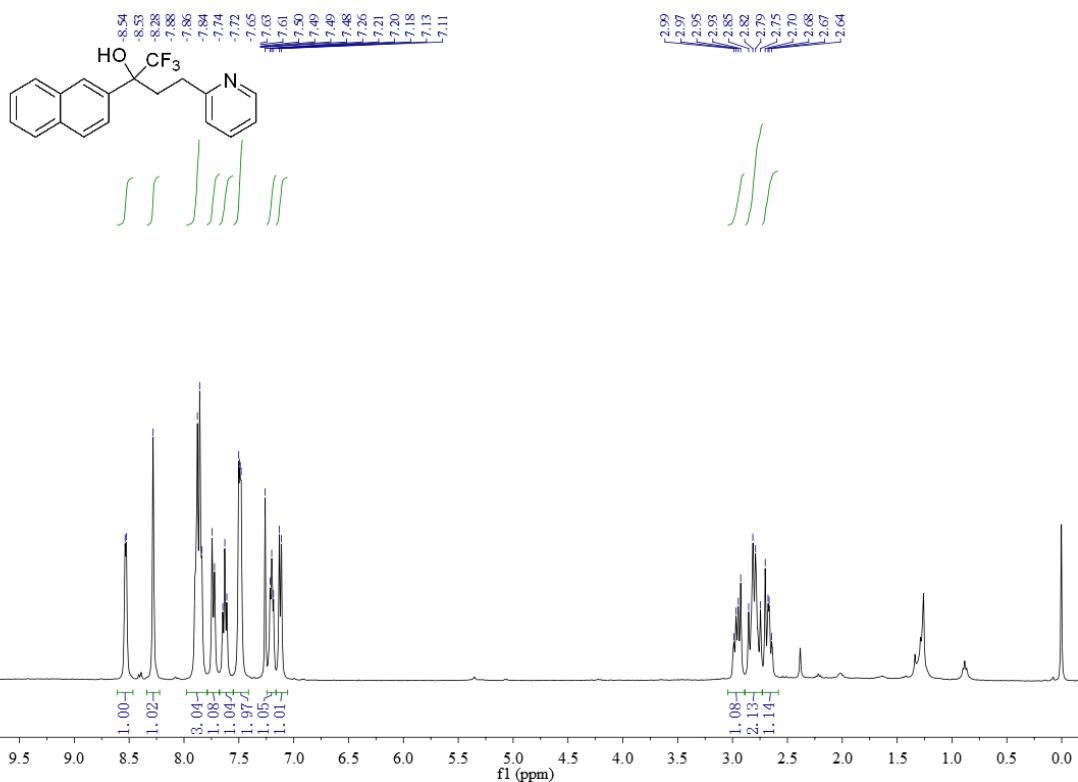
¹³C NMR of **3n**



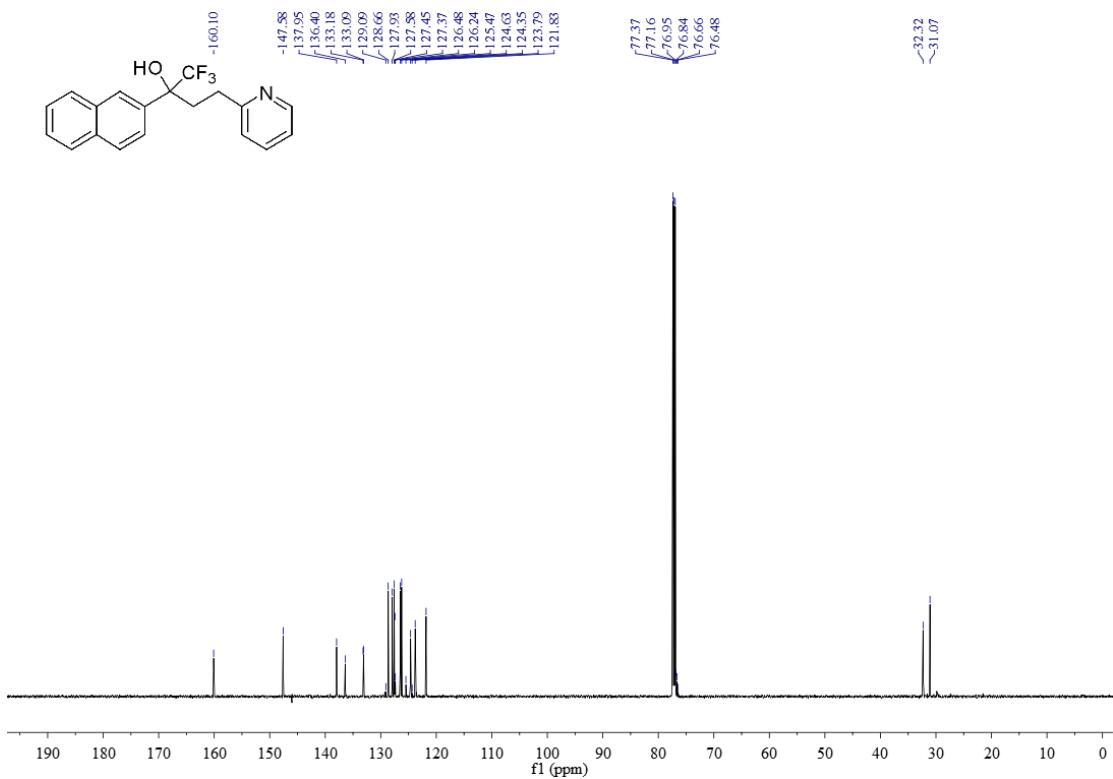
¹⁹F NMR of **3n**



¹H NMR of **3o**



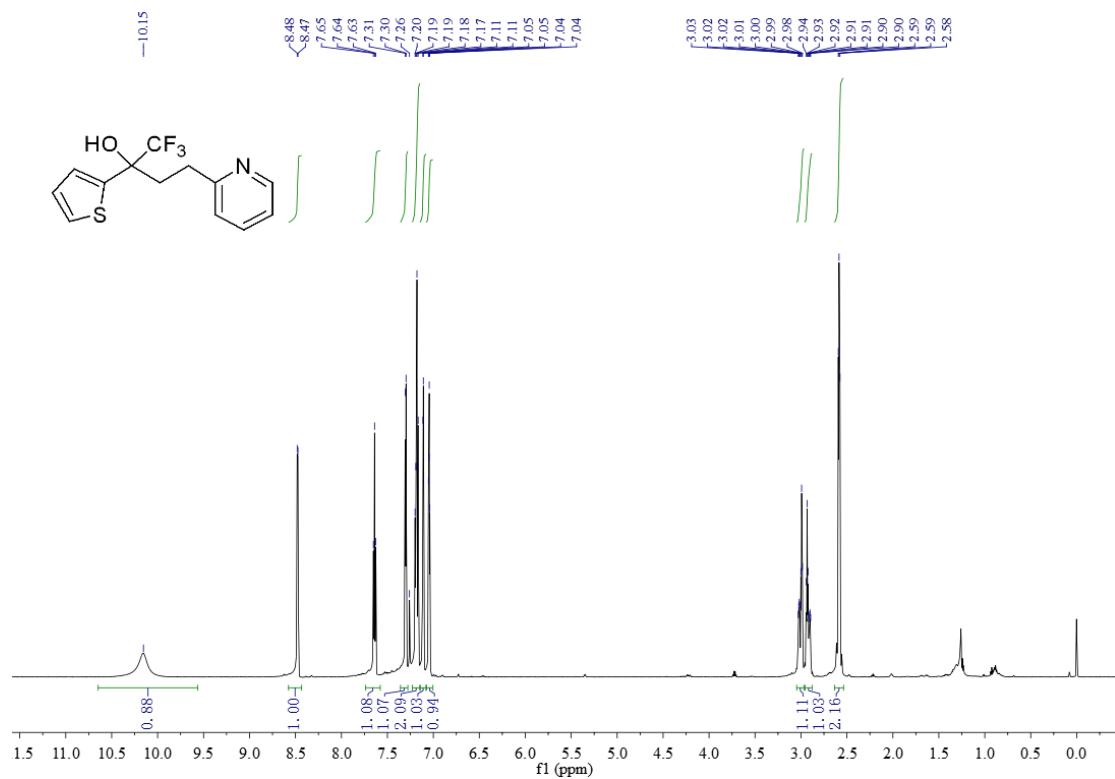
¹³C NMR of **3o**



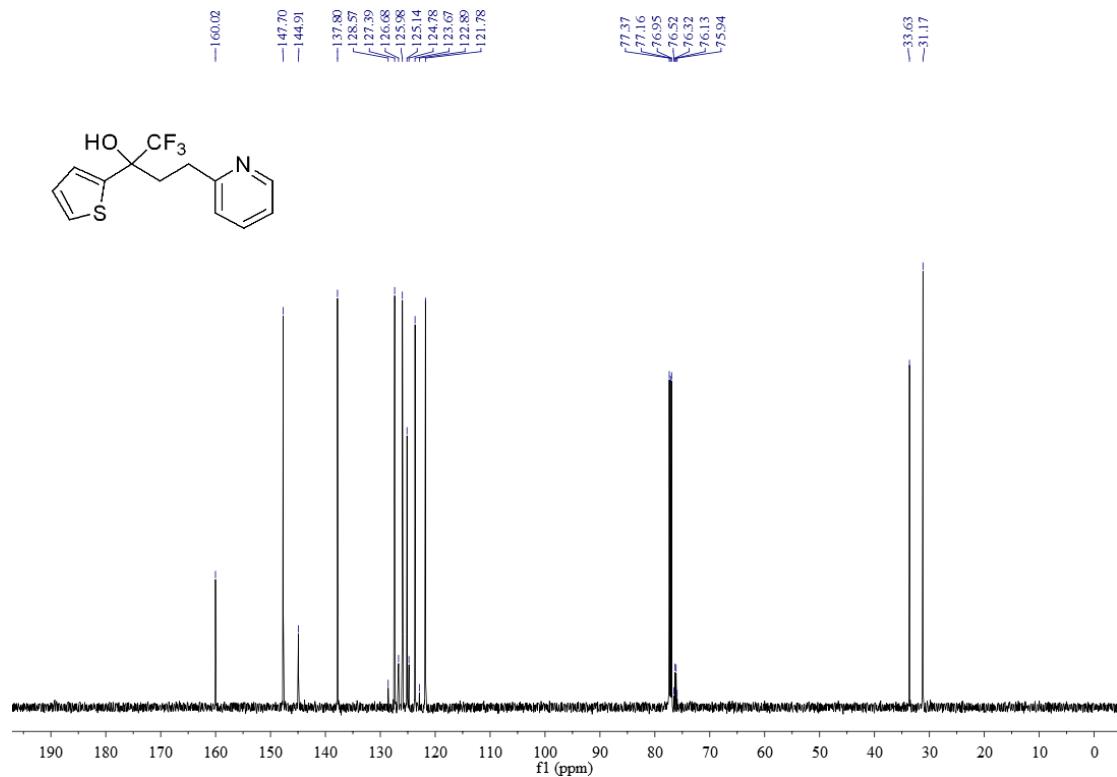
¹⁹F NMR of **3o**



¹H NMR of **3p**



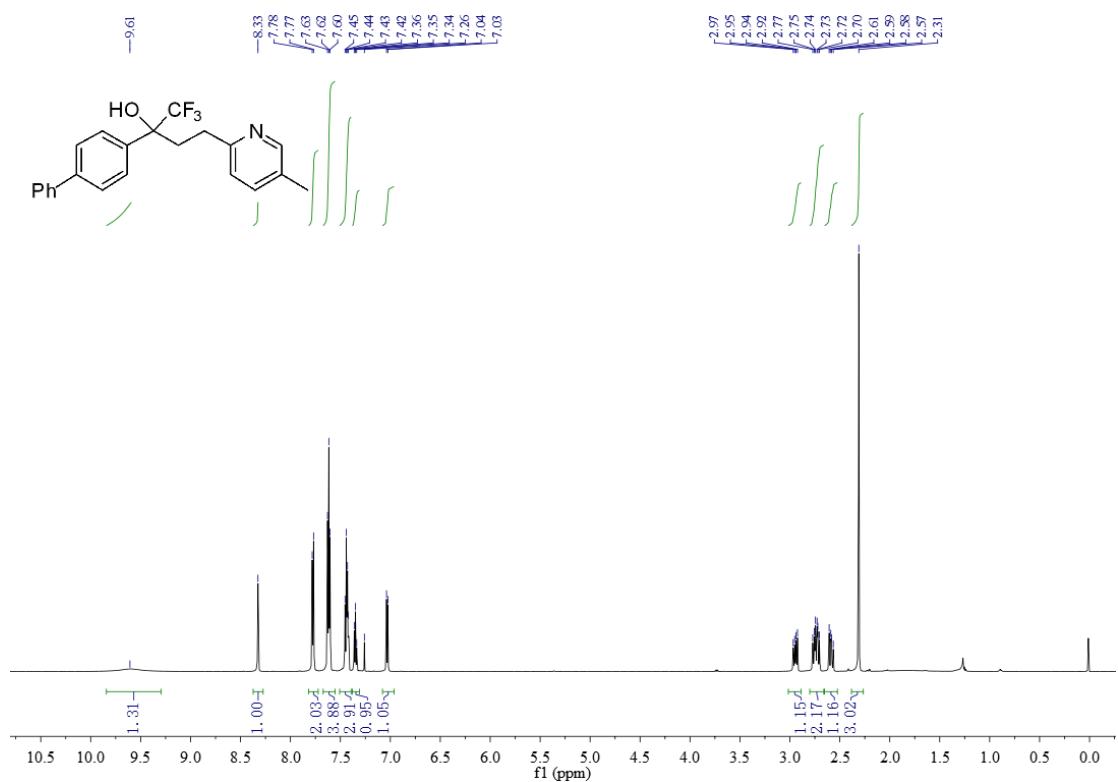
¹³C NMR of **3p**



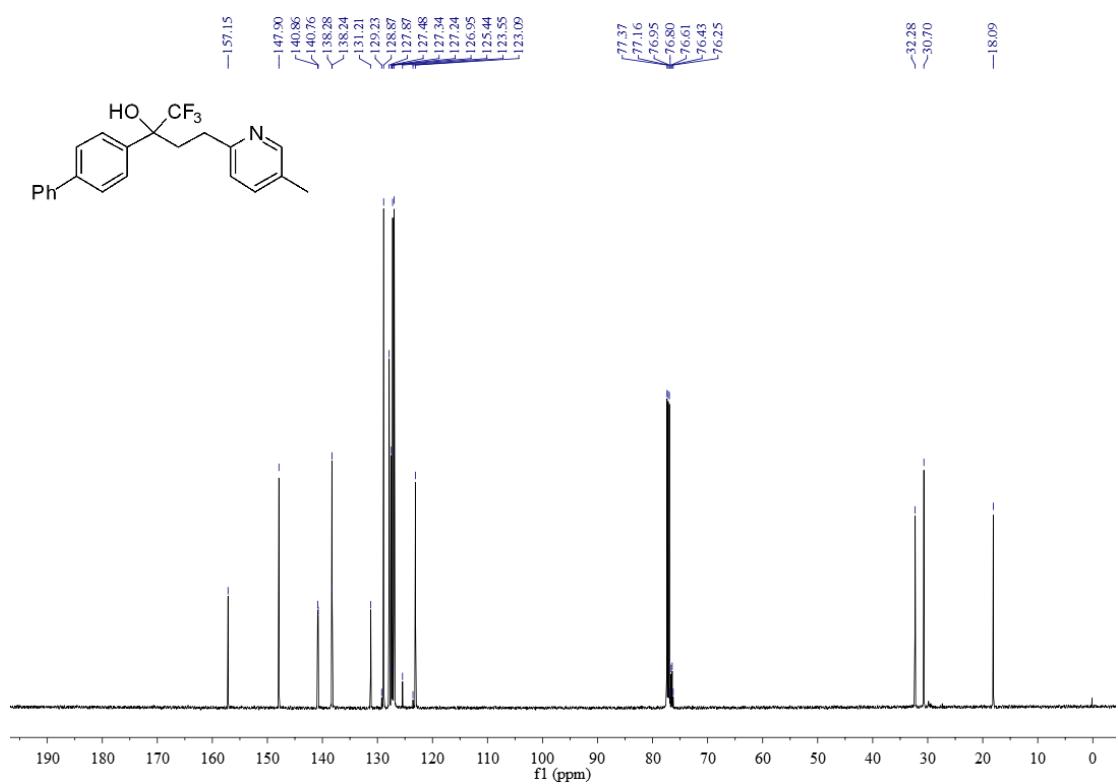
¹⁹F NMR of **3p**



¹H NMR of **3q**



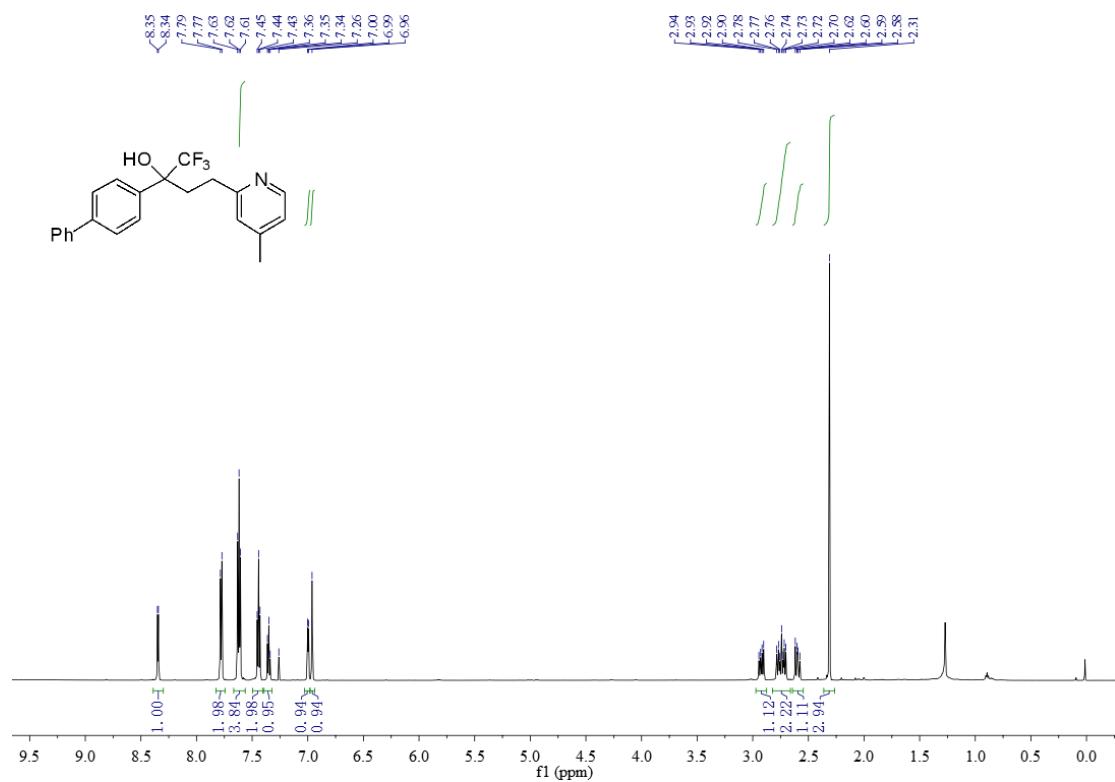
¹³C NMR of **3q**



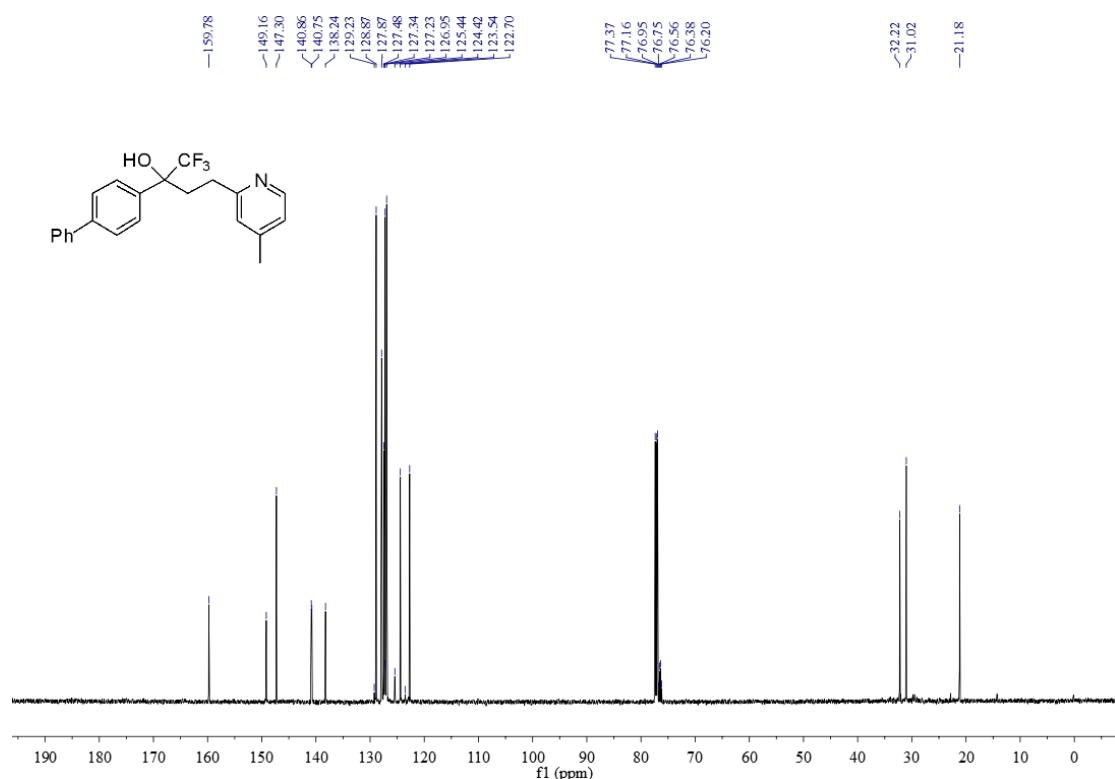
¹⁹F NMR of **3q**



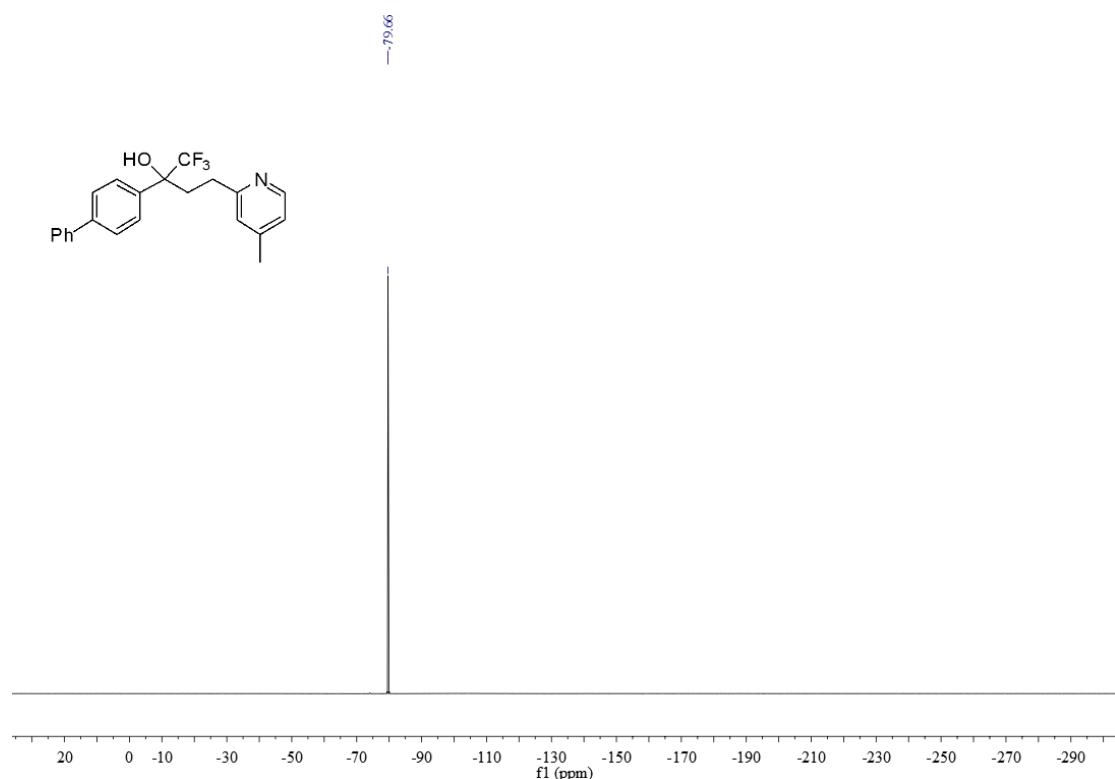
¹H NMR of **3r**



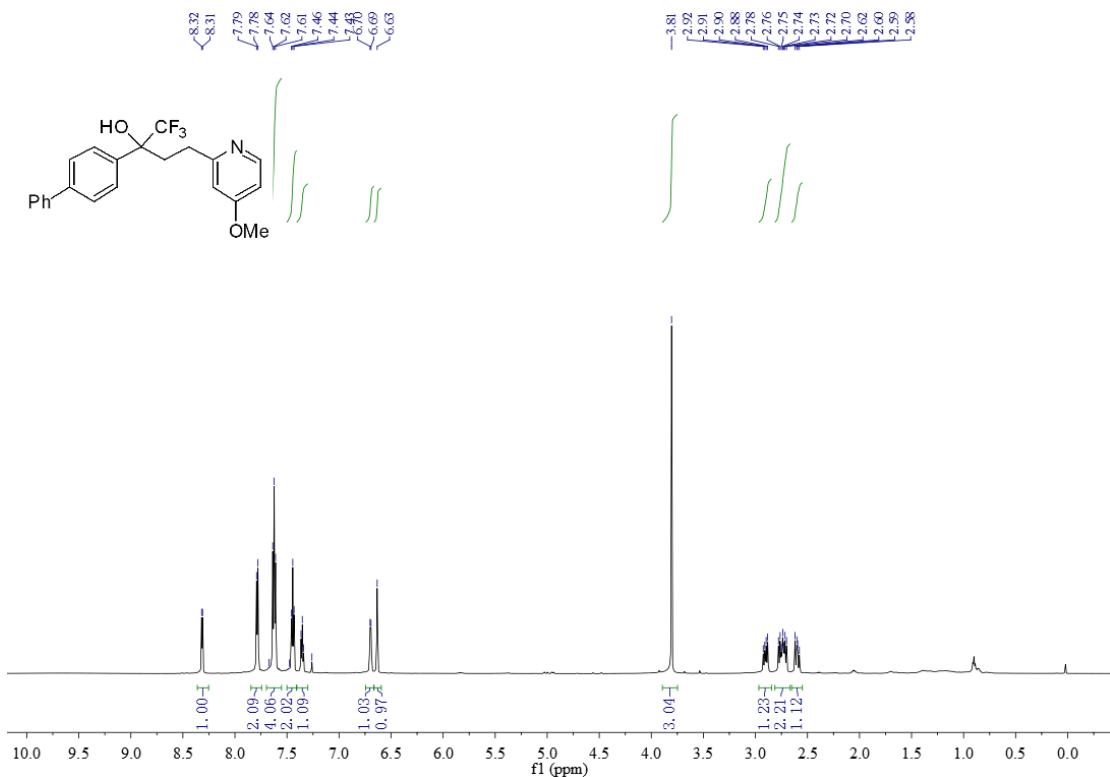
¹³C NMR of **3r**



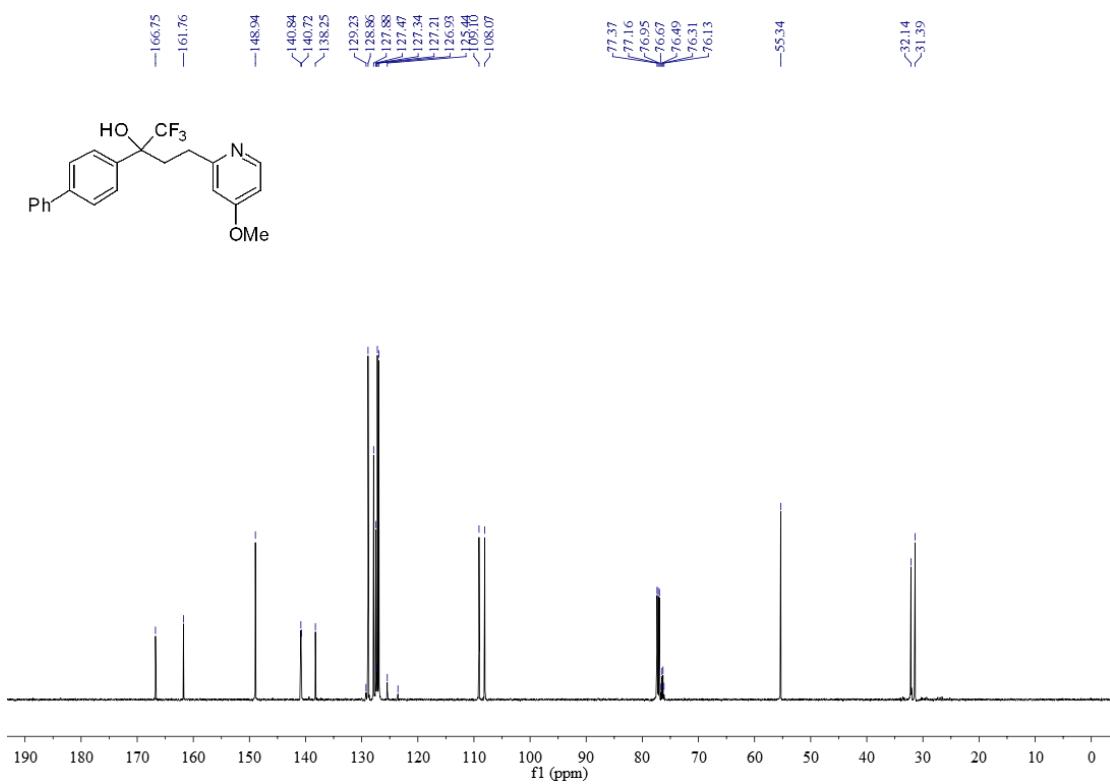
¹⁹F NMR of **3r**



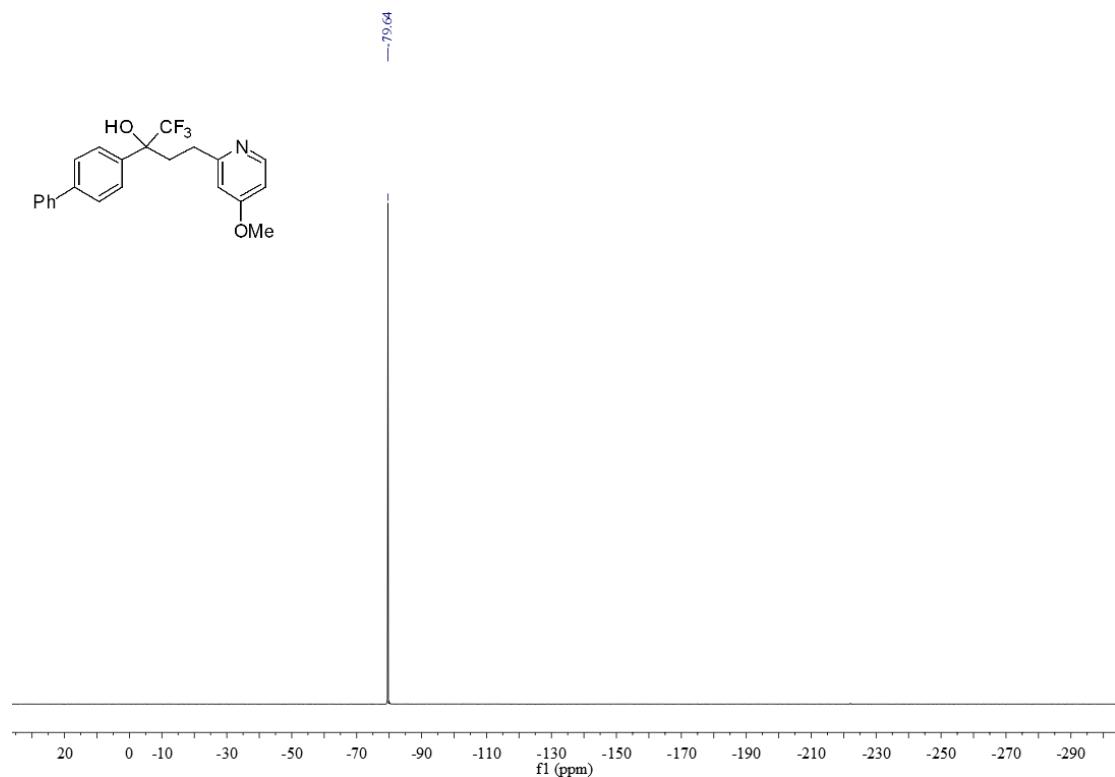
¹H NMR of **3s**



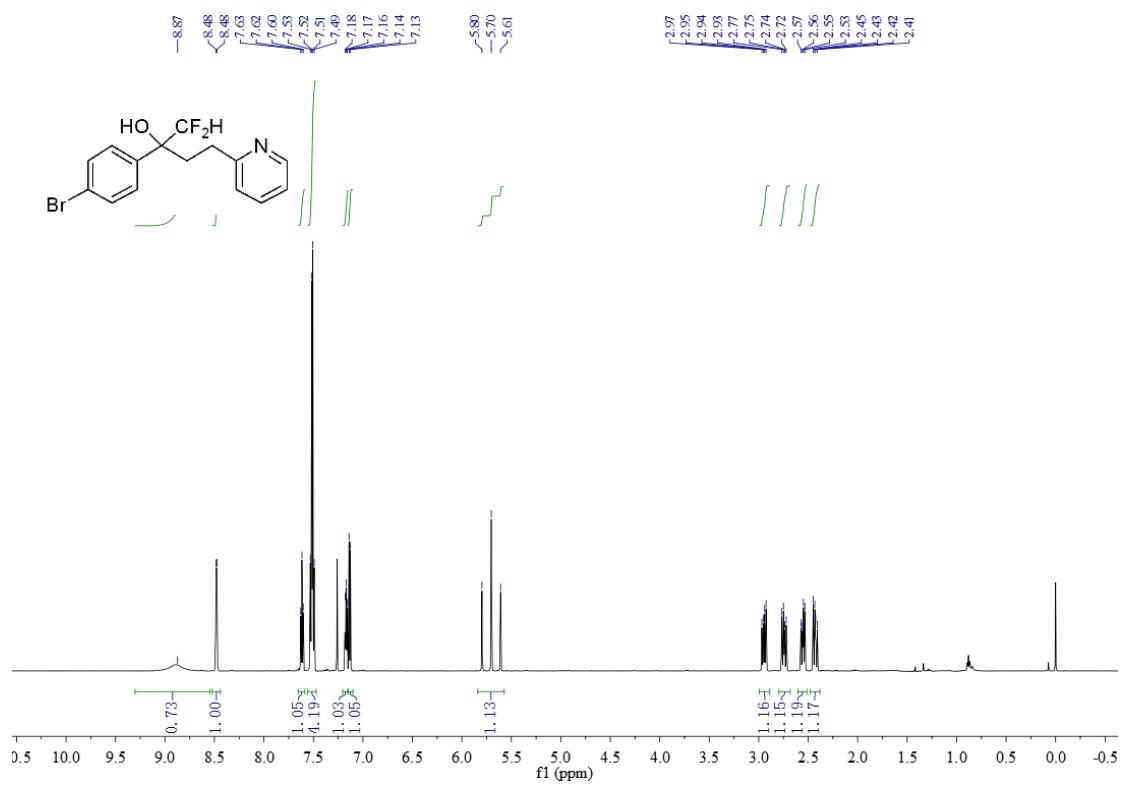
¹³C NMR of **3s**



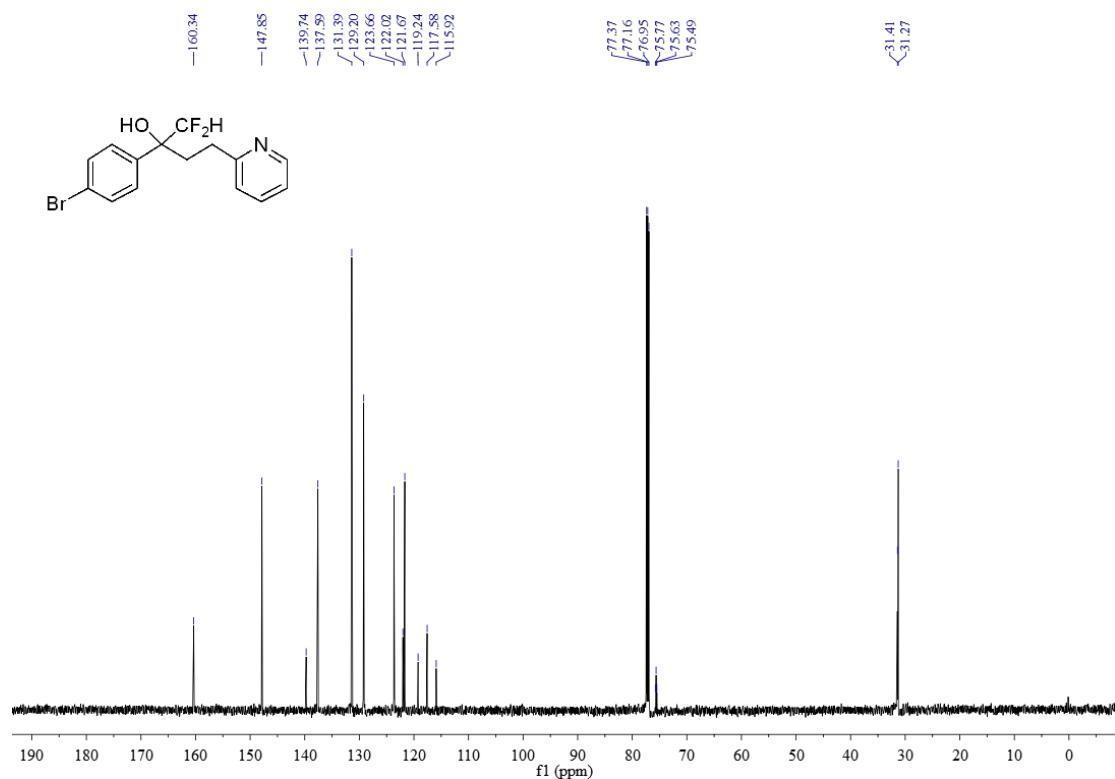
¹⁹F NMR of **3s**



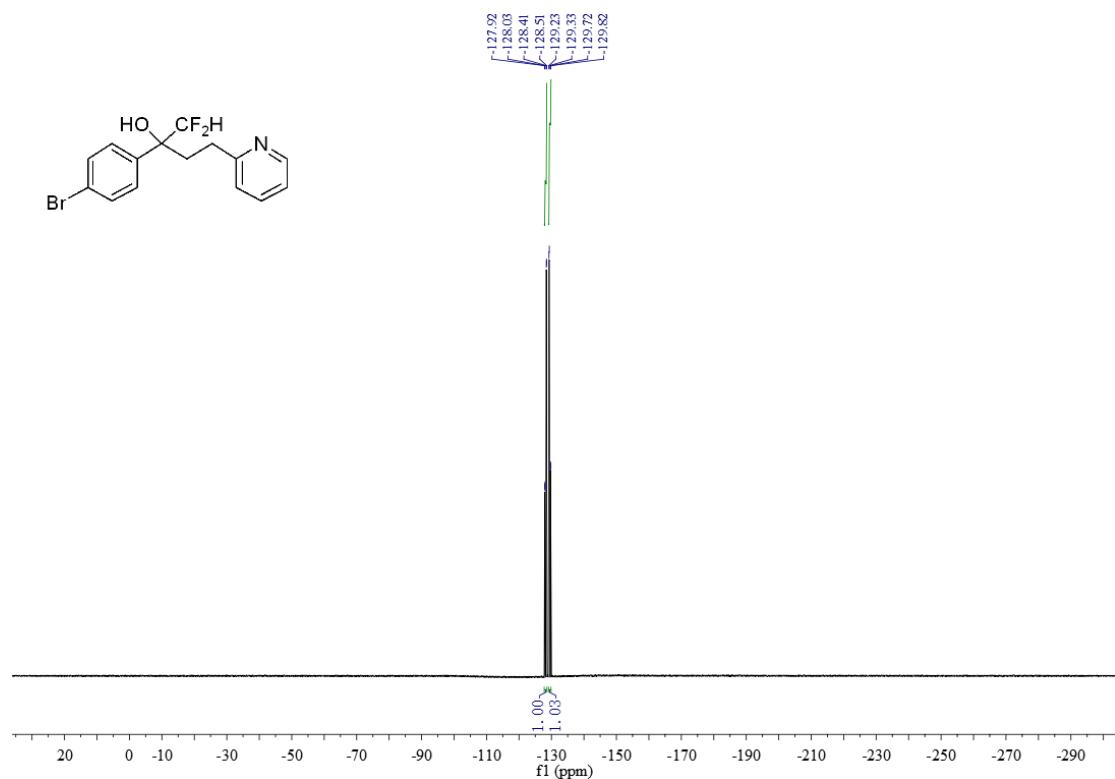
¹H NMR of **5a**



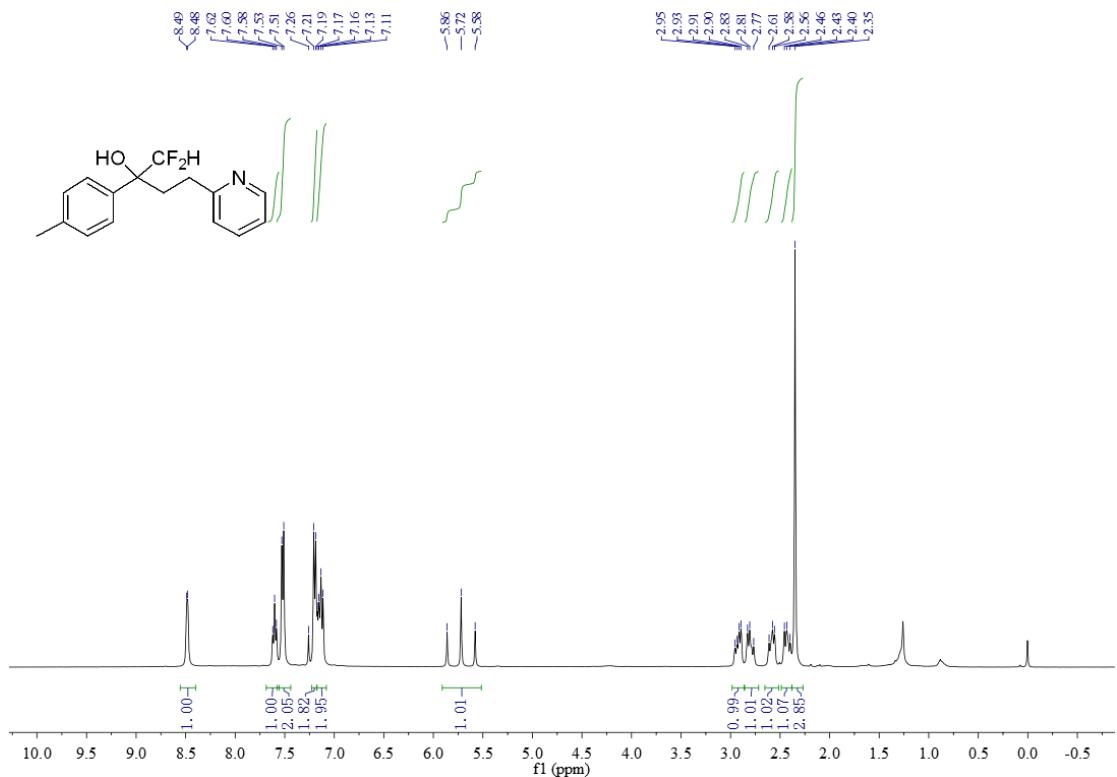
¹³C NMR of **5a**



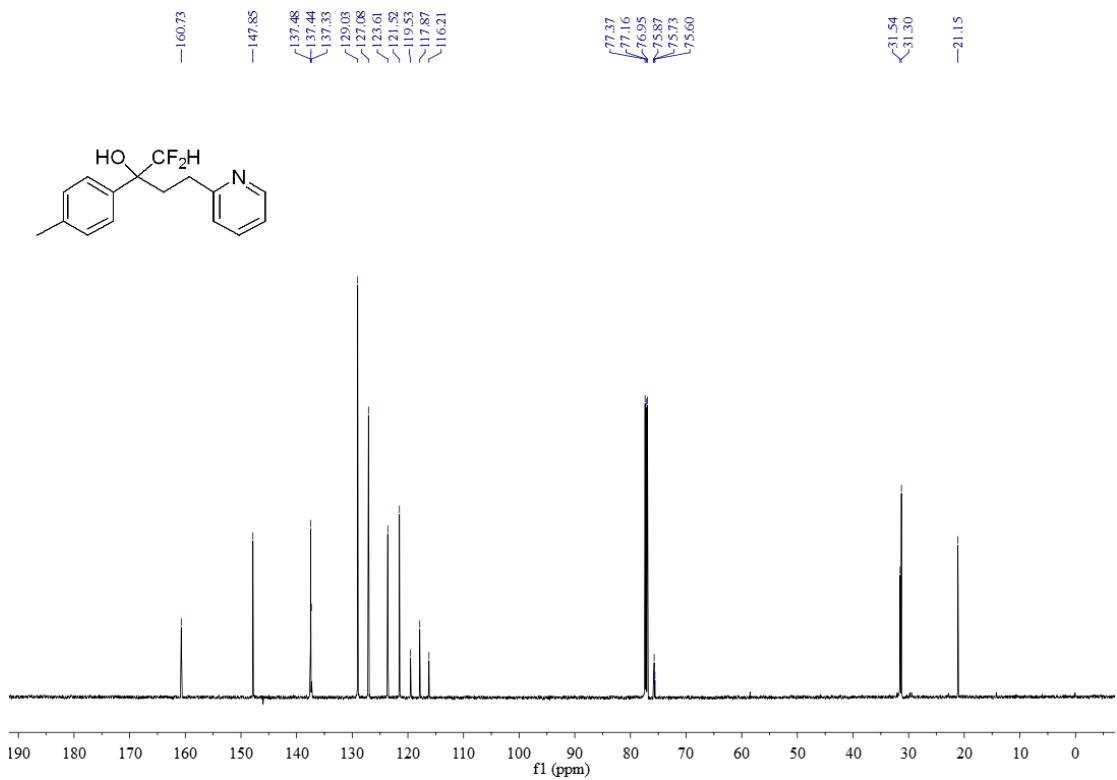
¹⁹F NMR of **5a**



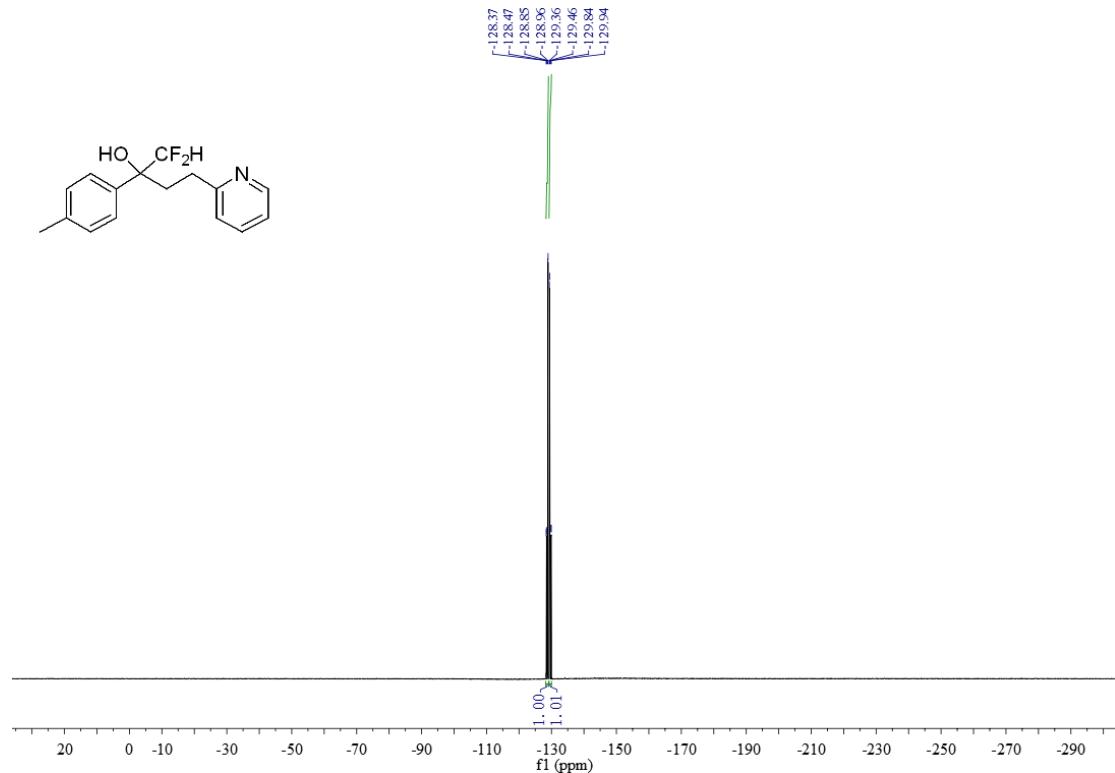
¹H NMR of **5b**



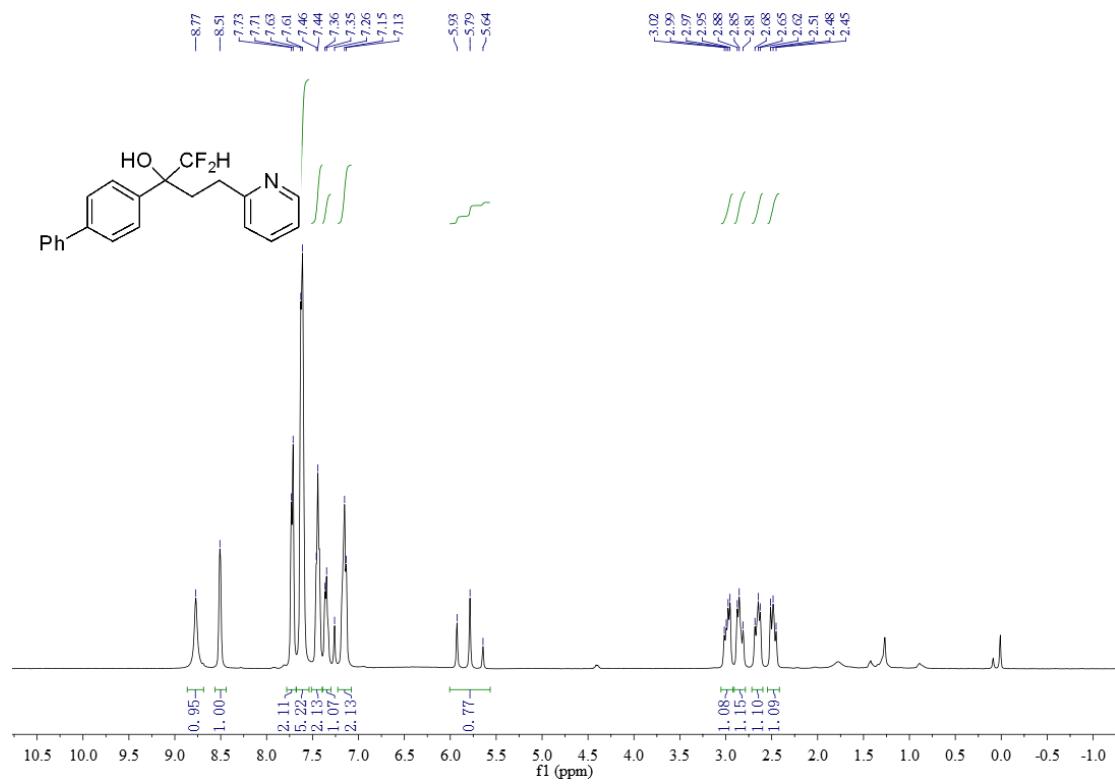
¹³C NMR of **5b**



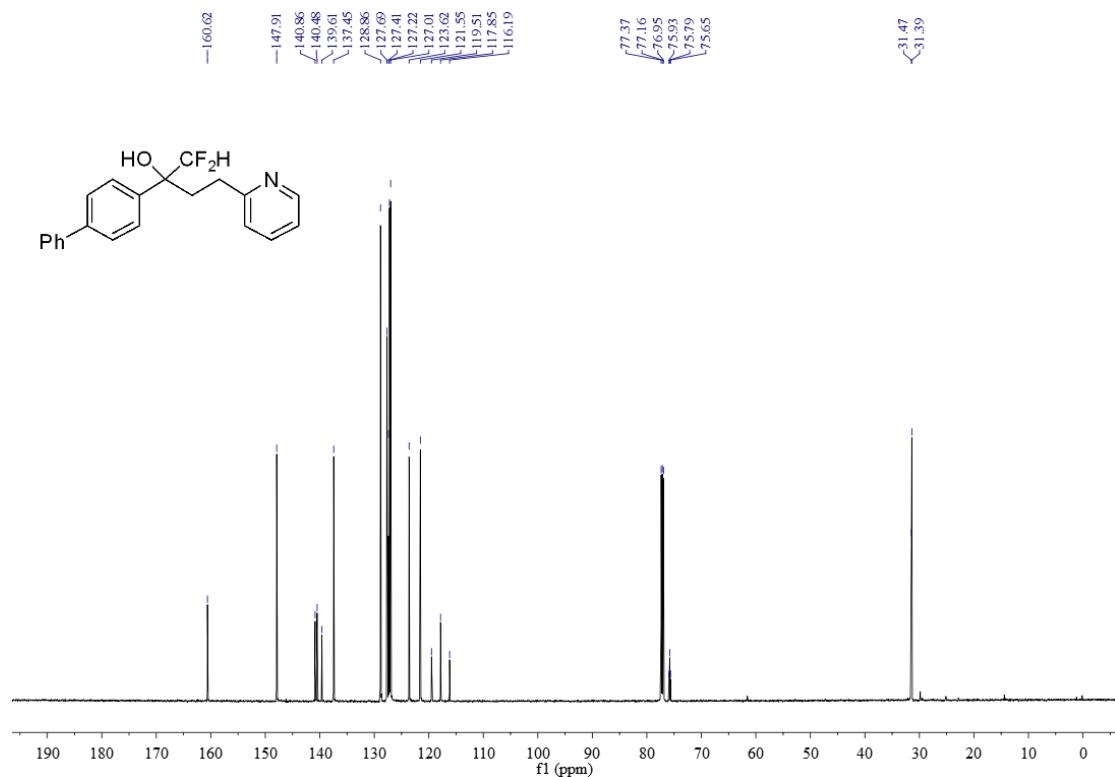
¹⁹F NMR of **5b**



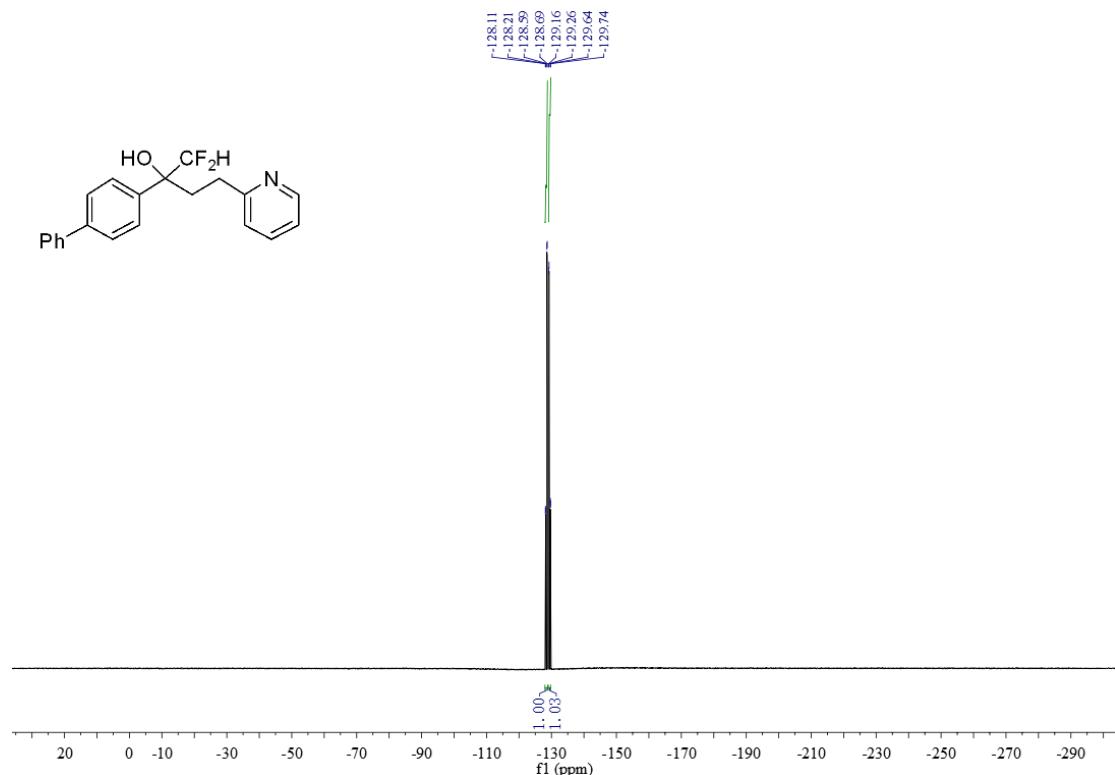
¹H NMR of **5c**



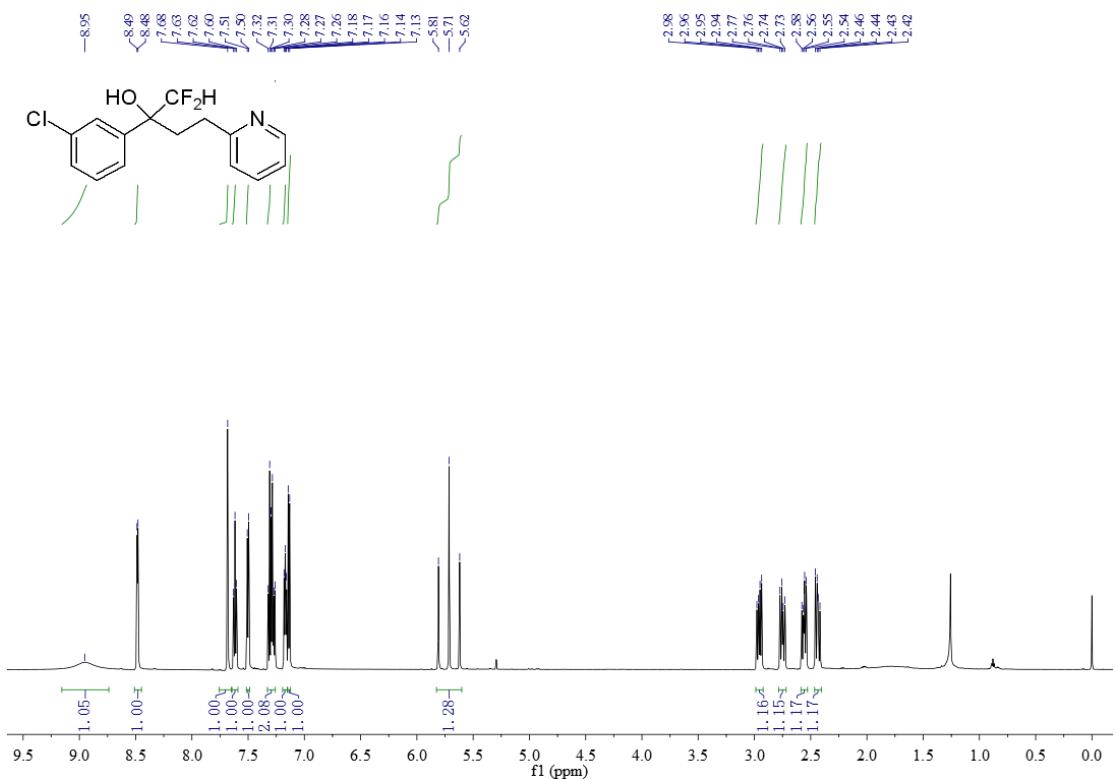
¹³C NMR of **5c**



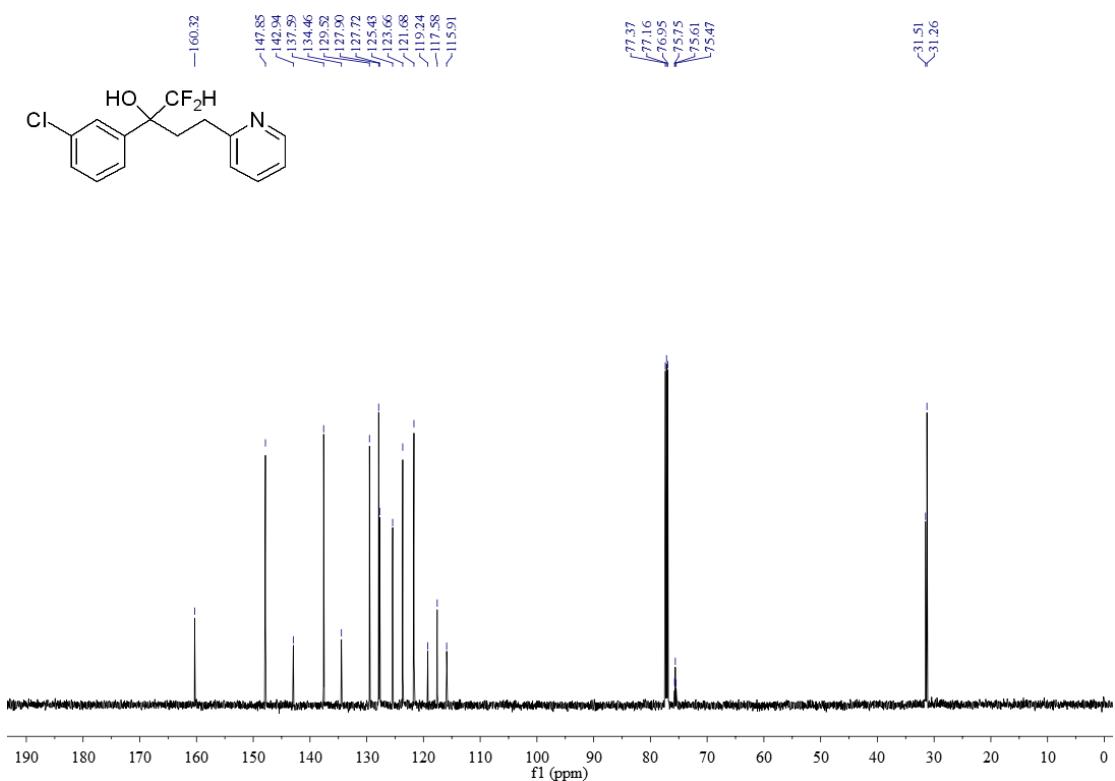
¹⁹F NMR of **5c**



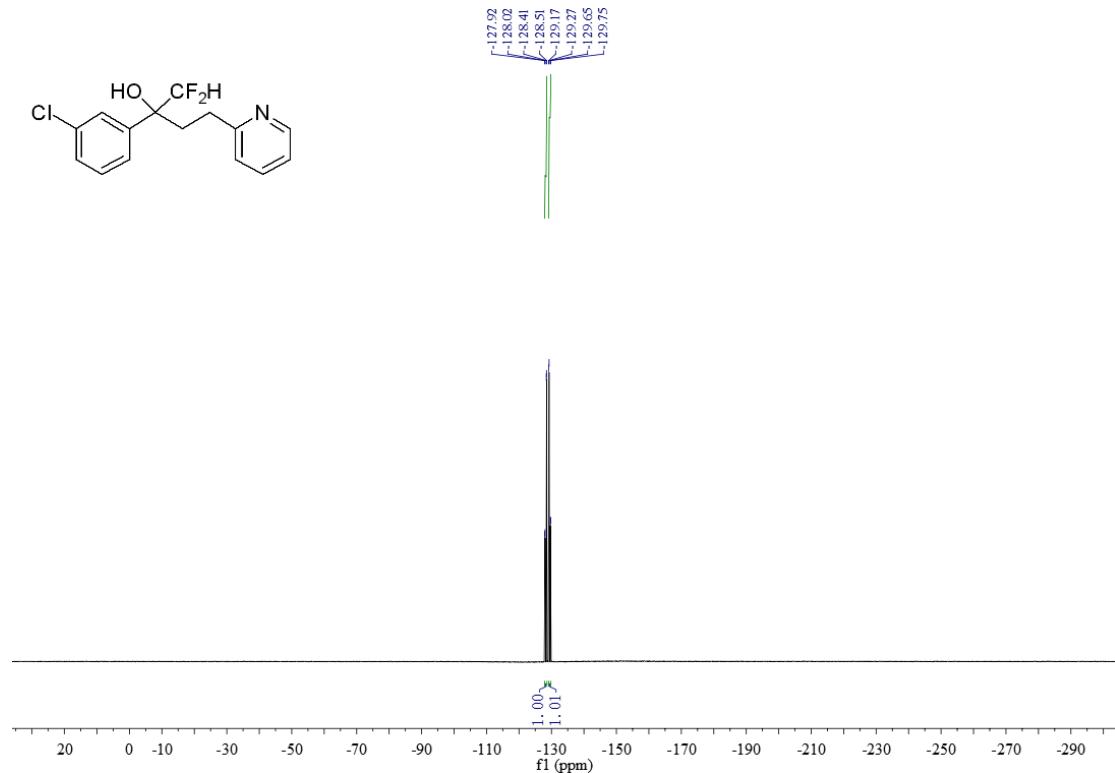
¹H NMR of **5d**



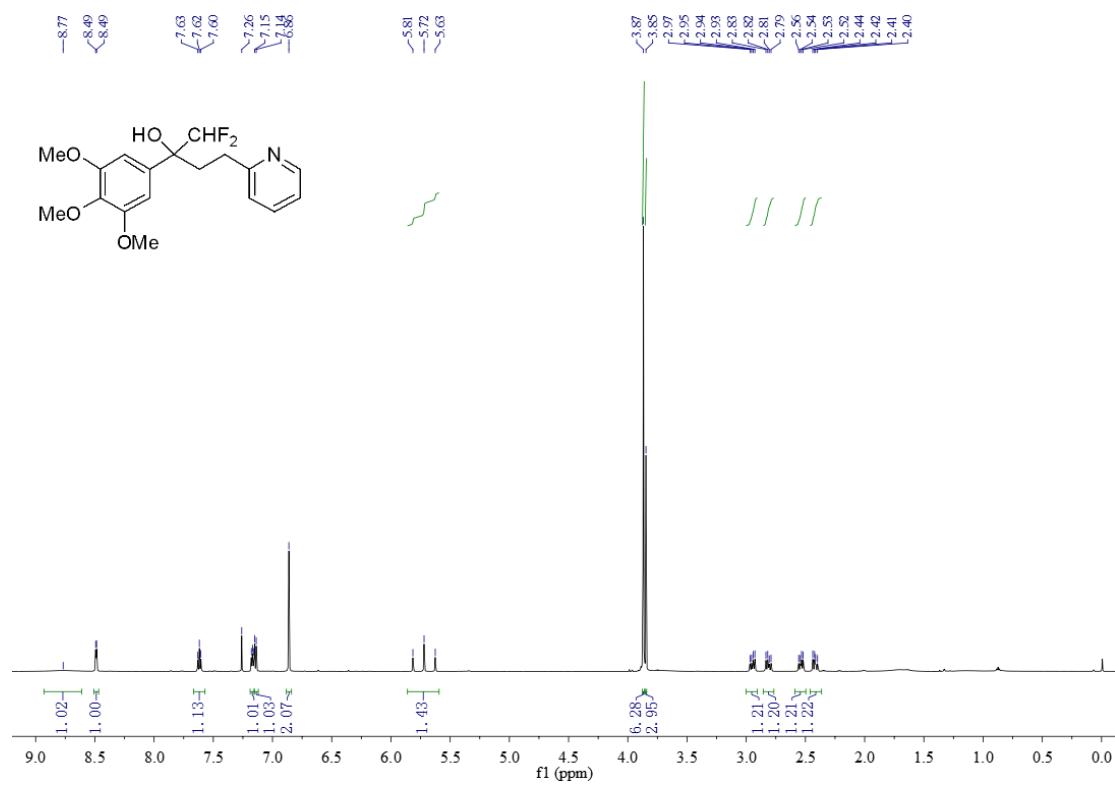
¹³C NMR of **5d**



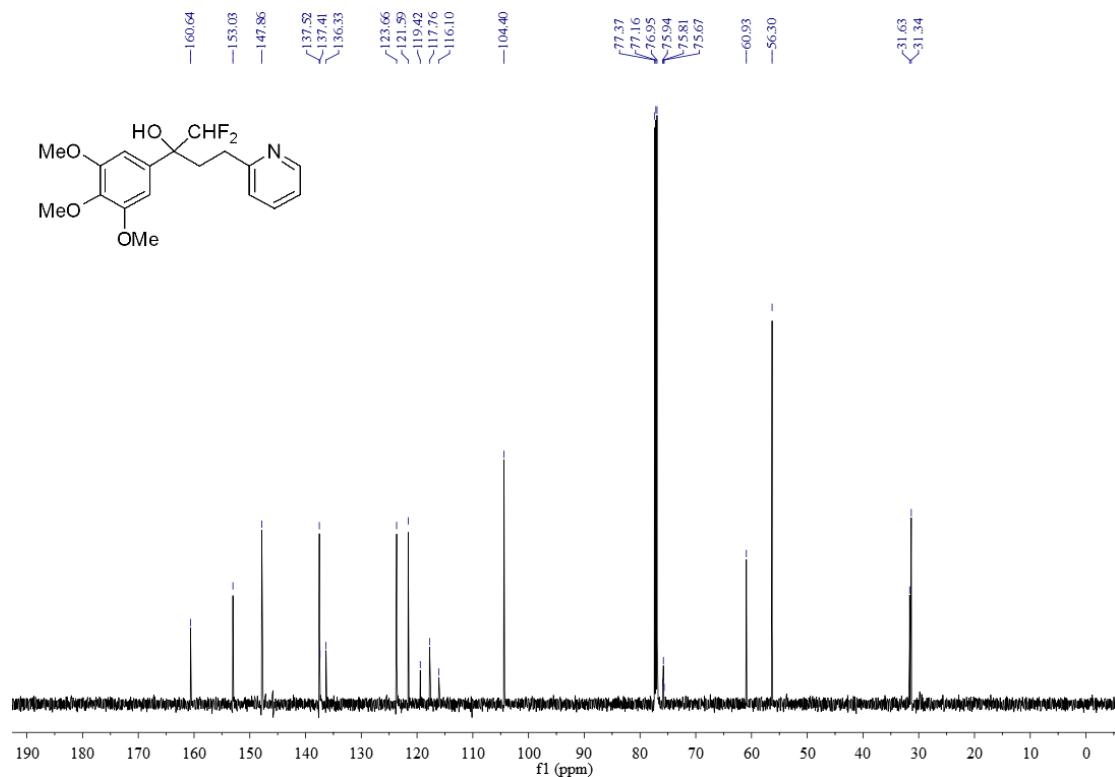
¹⁹F NMR of **5d**



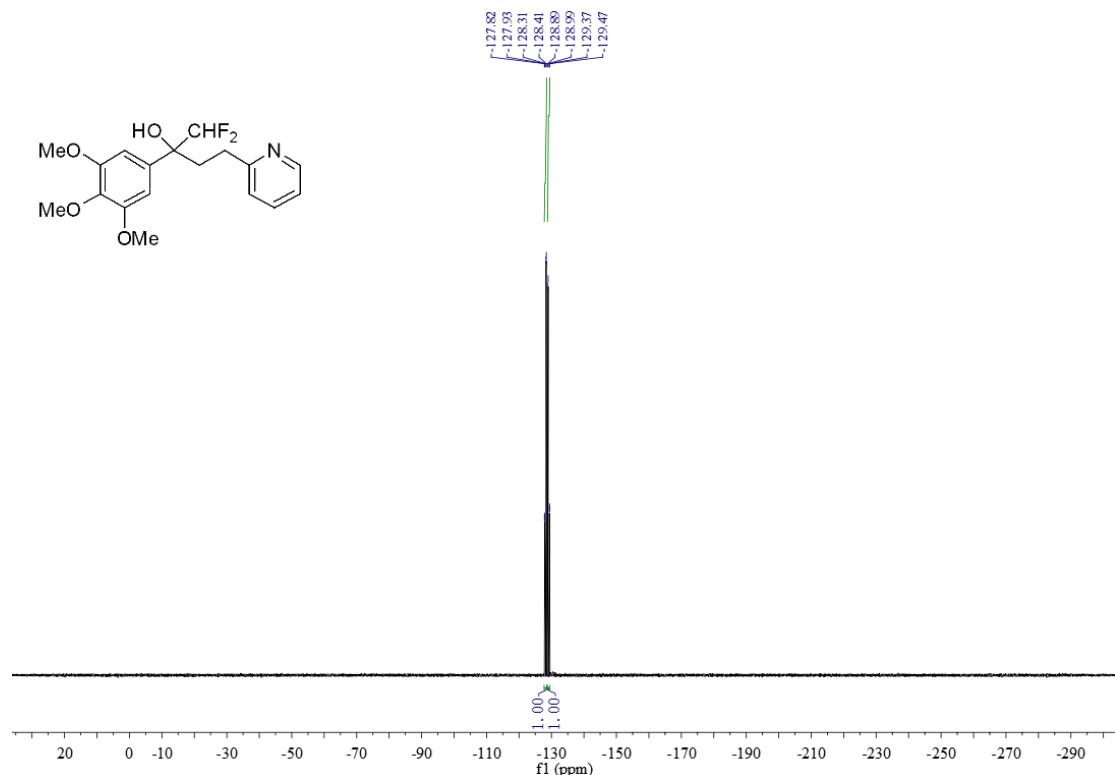
¹H NMR of **5e**



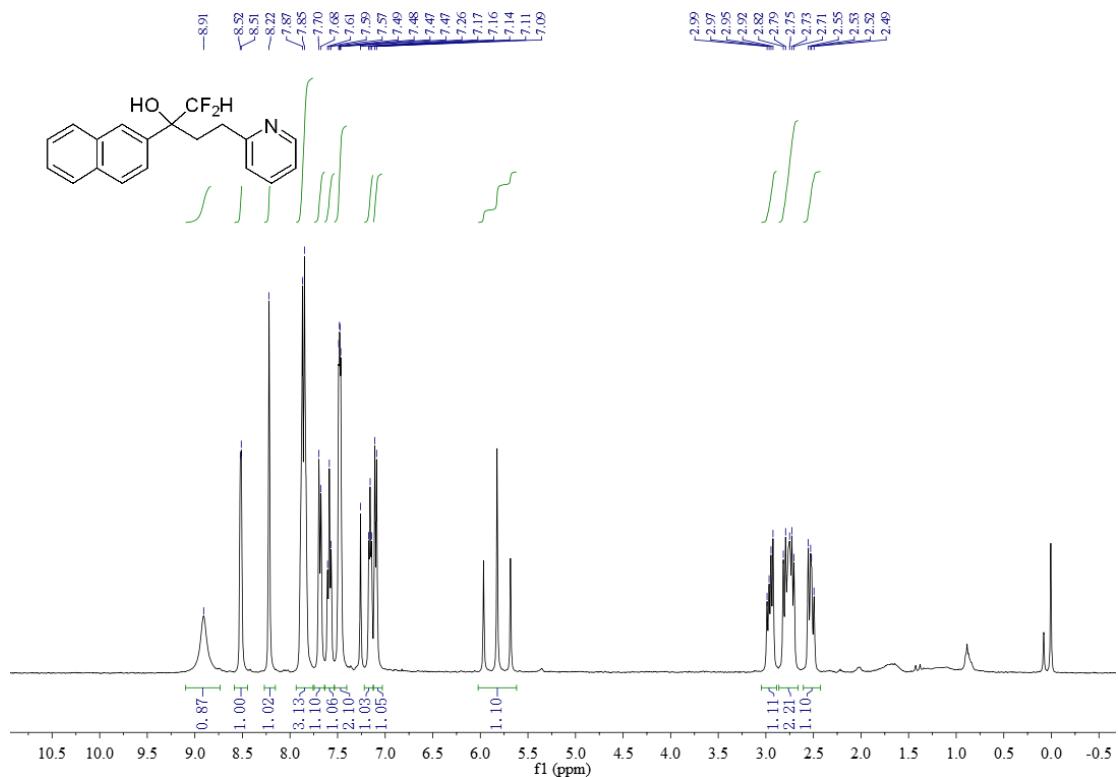
¹³C NMR of **5e**



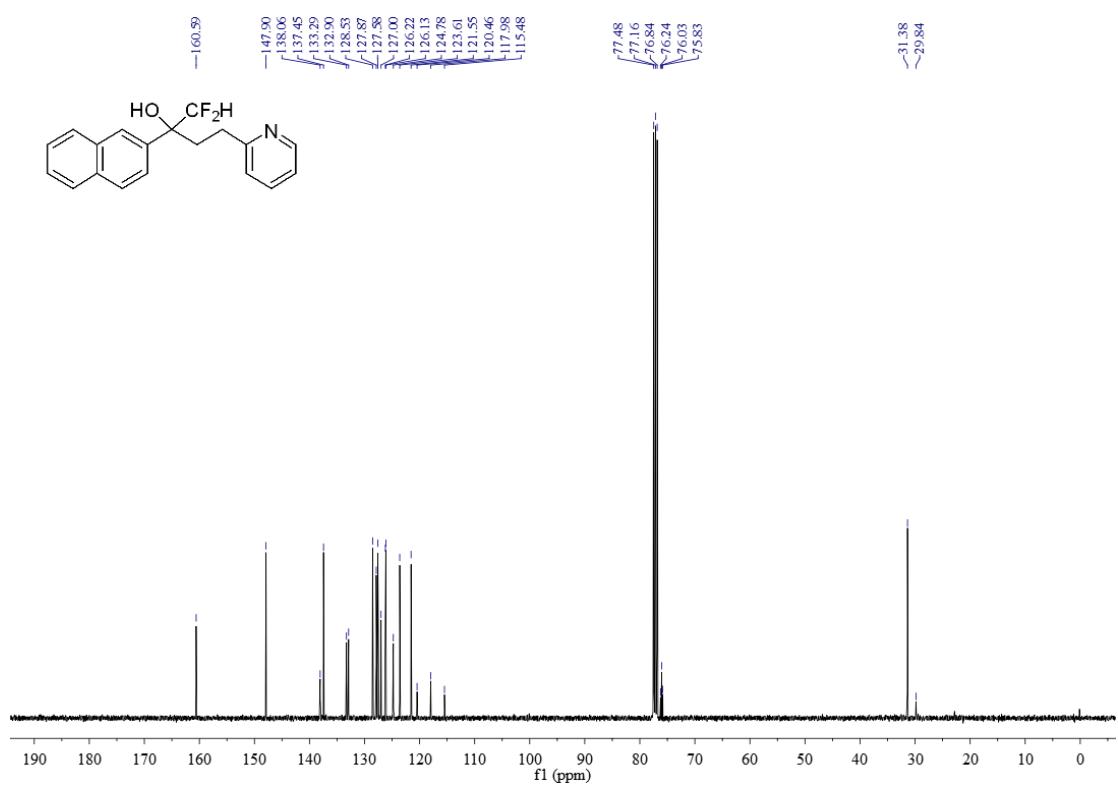
¹⁹F NMR of **5e**



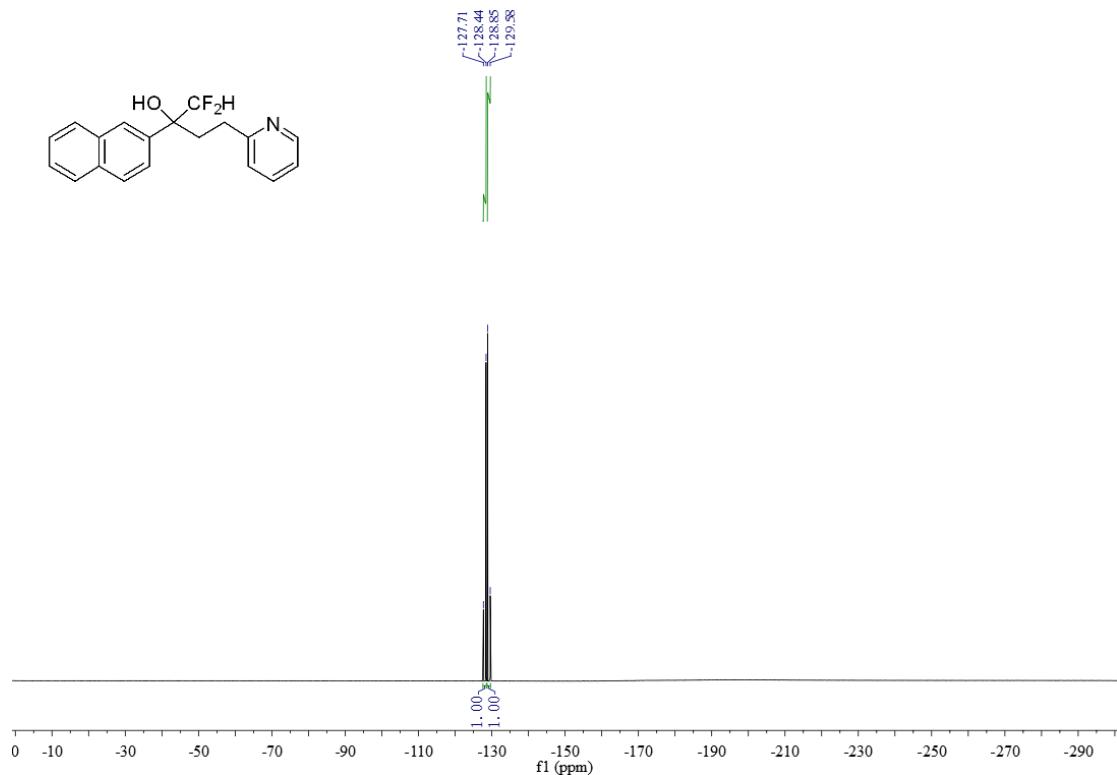
¹H NMR of **5f**



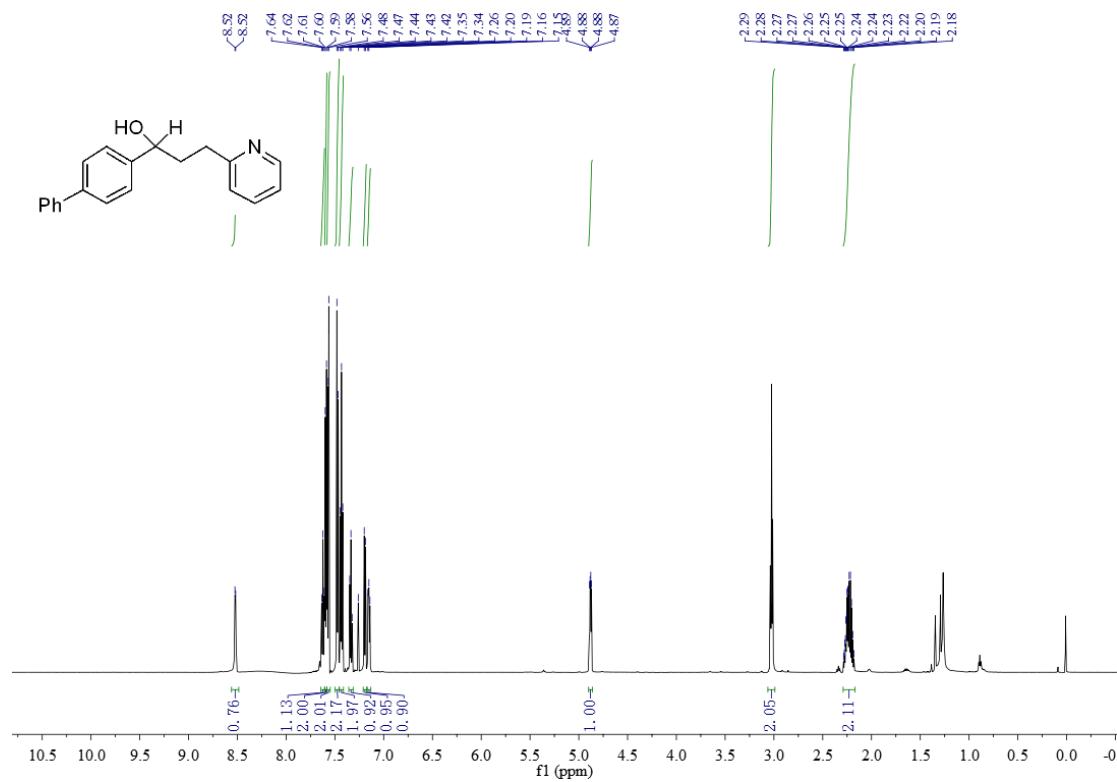
¹³C NMR of **5f**



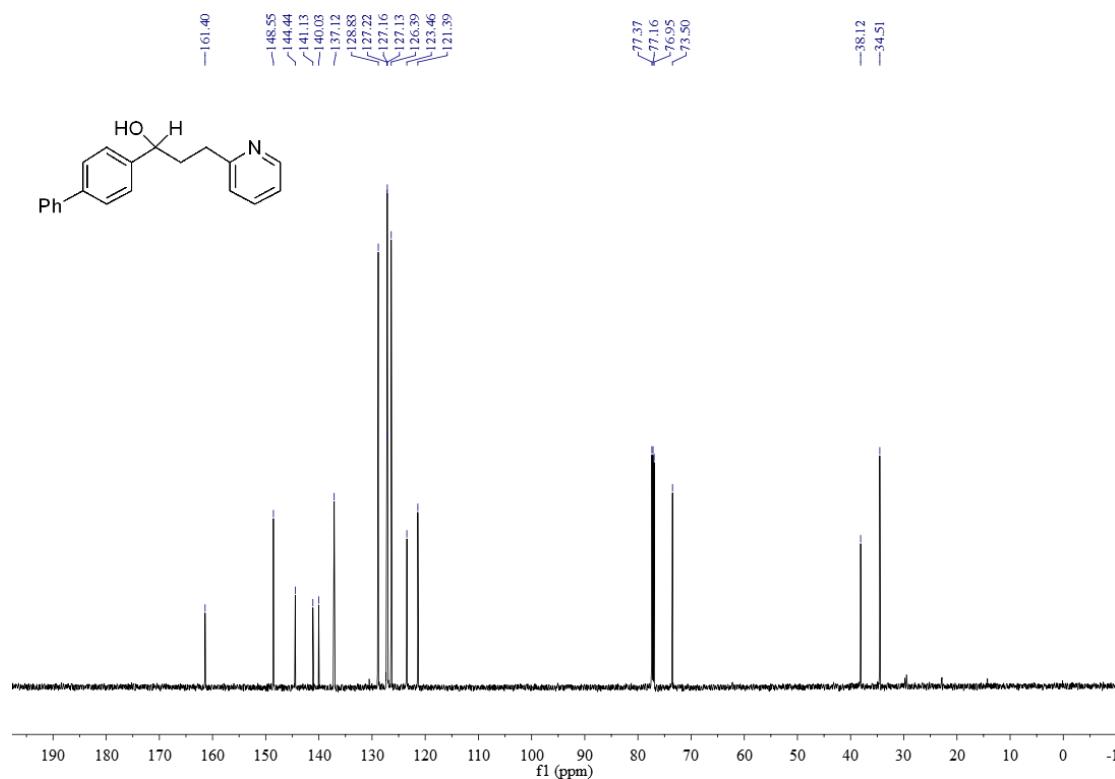
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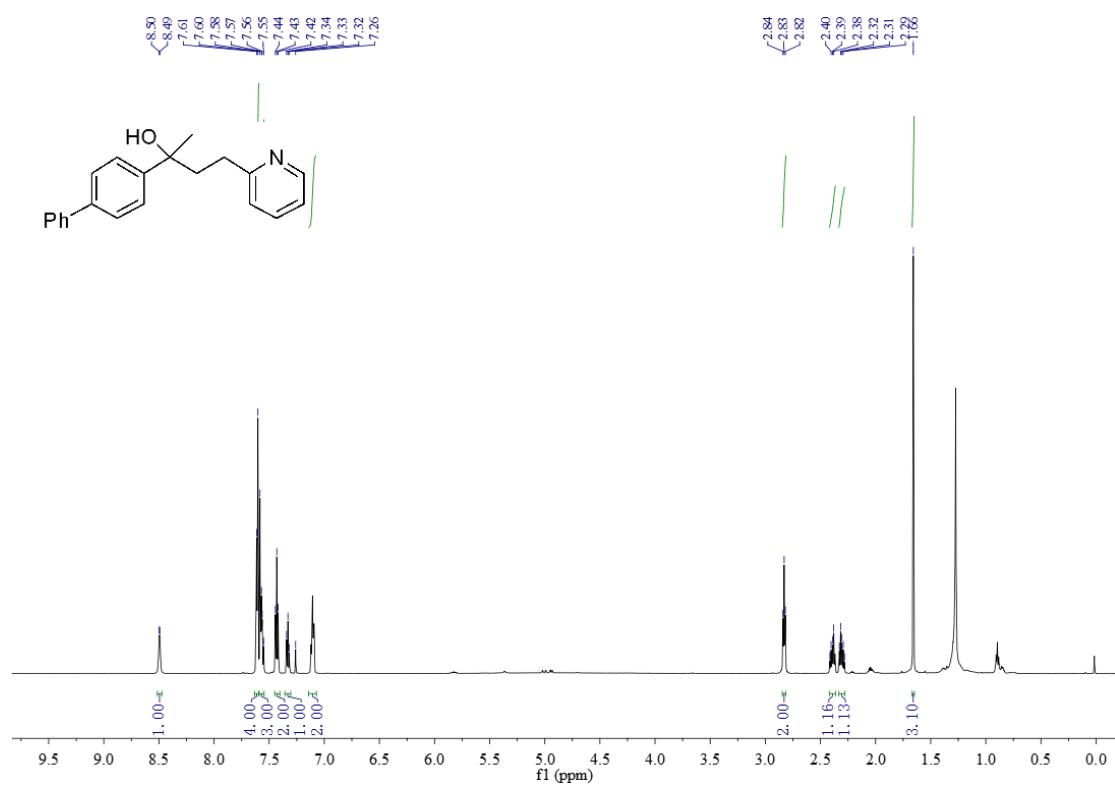
¹H NMR of **7a**



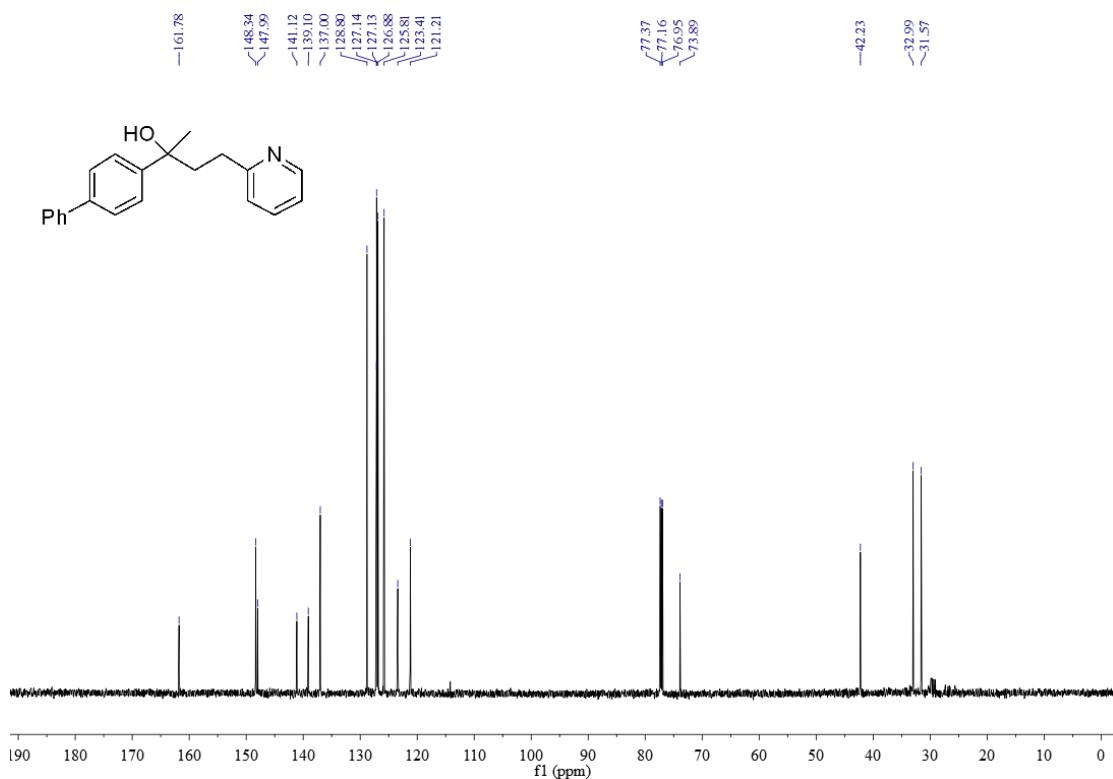
¹³C NMR of **7a**



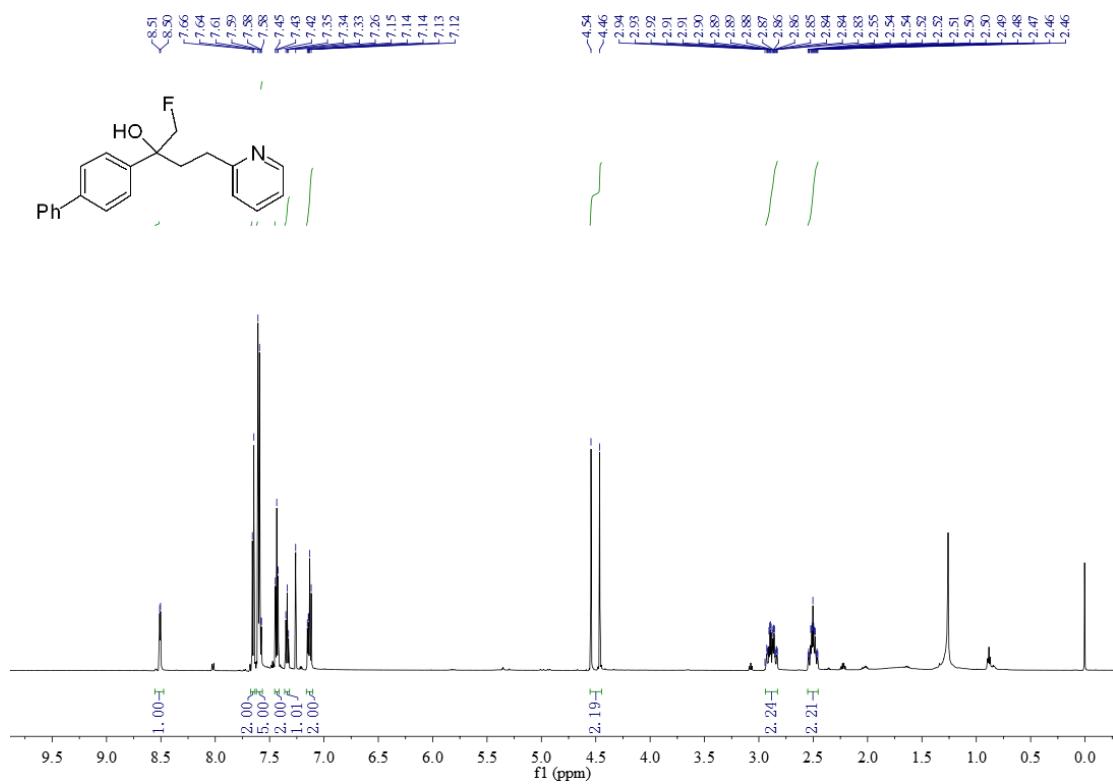
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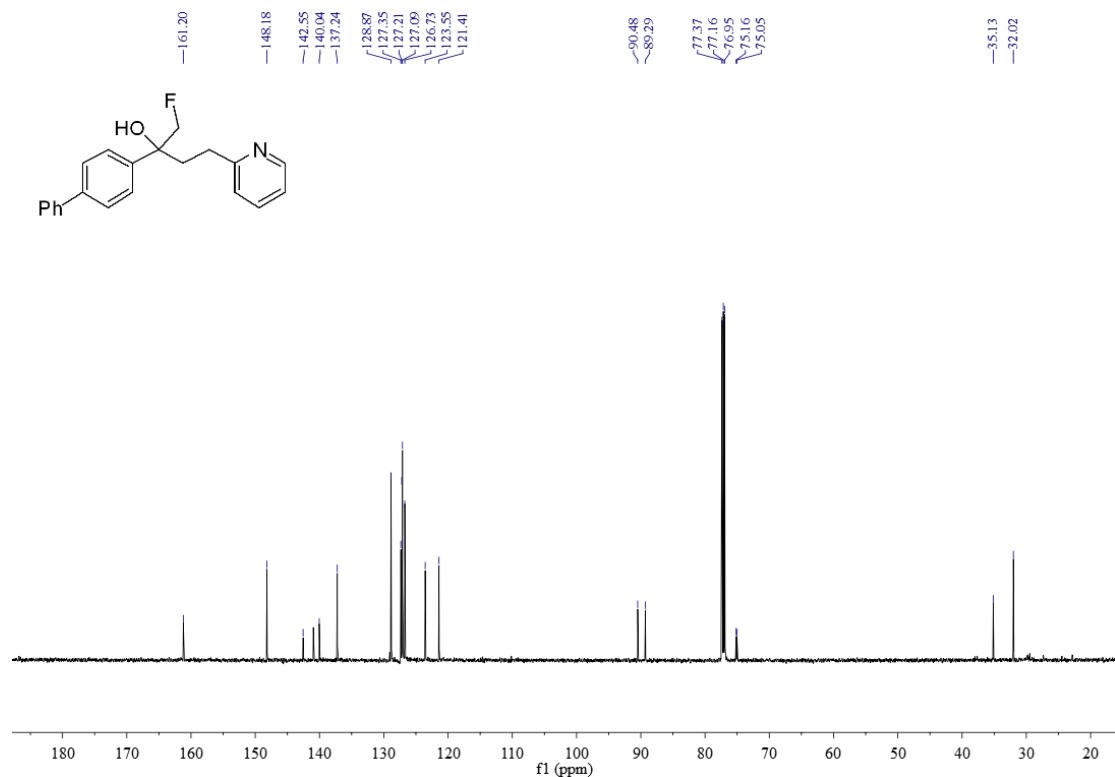
¹³C NMR of **7b**



¹H NMR of **7c**



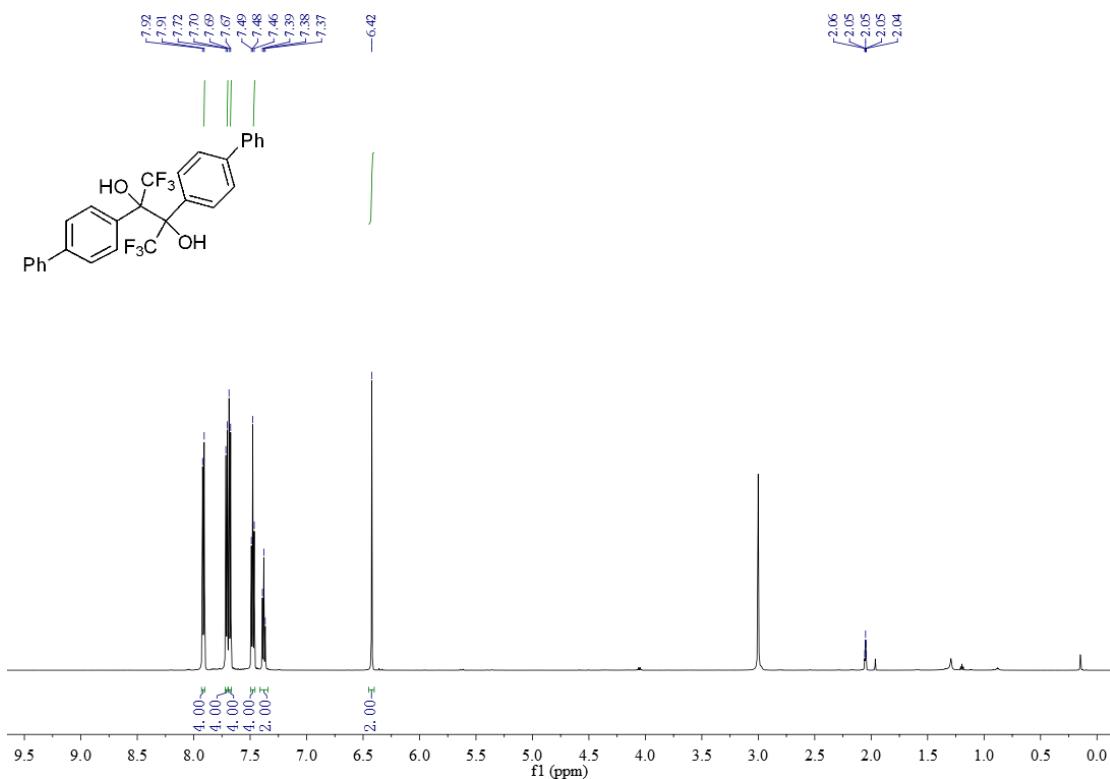
¹³C NMR of **7c**



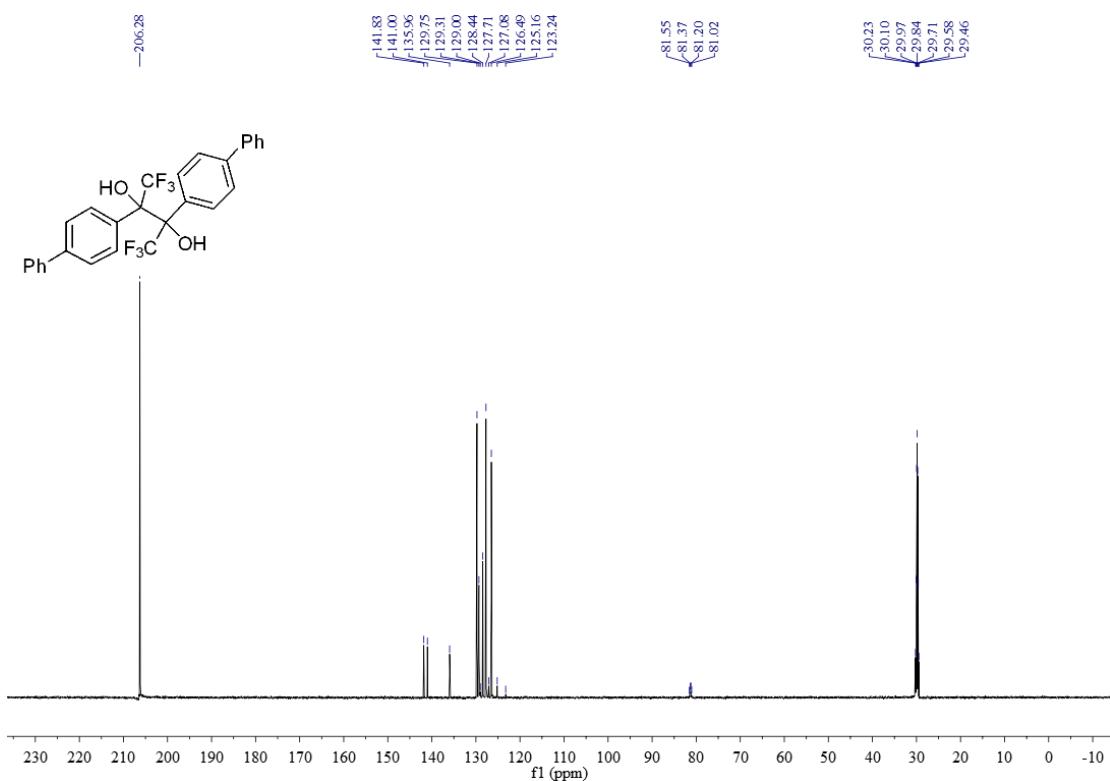
¹⁹F NMR of **7c**



¹H NMR of **9**



¹³C NMR of **9**



¹⁹F NMR of **9**

