

Supporting Information

Organophotocatalytic acyltrifluoromethylation of unactivated alkenes enabled the efficient synthesis of CF₃-substituted chroman-4-one

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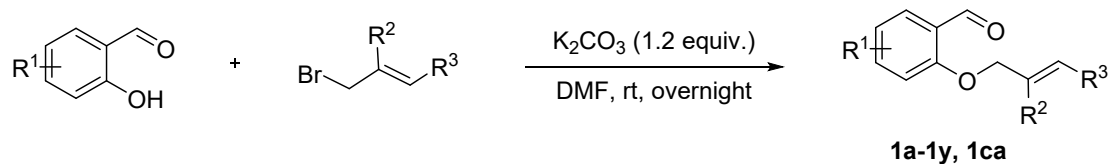
(A) General Information

All reactions and manipulations which are sensitive to moisture or air were performed under inert atmosphere of argon. All chemicals were purchased from J&K, Acros and Aldrich, and were used as received. Anhydrous THF, DMSO, 1,4-dioxane, DMF and MeCN were freshly distilled from calcium hydride. Anhydrous MeOH freshly distilled from magnesium. ^1H NMR, ^{13}C NMR spectra were recorded on a Bruker AVANCE 400 and chemical shifts are reported in δ (ppm) referenced to residual undeuterated solvent signal for ^1H NMR (7.26 ppm) and ^{13}C NMR (77 ppm). The following abbreviations were used to designate chemical shift multiplicities: s = singlet, d = doublet, t = triplet, q = quartet, m = multiplet. All first-order splitting patterns were assigned on the basis of the appearance of the multiplet. HRMS spectra were recorded on a Waters Acquity UPLC/Xevo TQD-MS-MS quadrupole mass spectrometer. The light source for the photocatalytic reaction is manufactured by GeAo chemistry with a power of 24 W blue LED (450–465 nm). A fan was used to maintain the reaction temperature at room temperature (about 25–30 °C). The reactions were carried out in a borosilicate glass vessel and the distance from the light source to the irradiation vessel is about 1 cm.

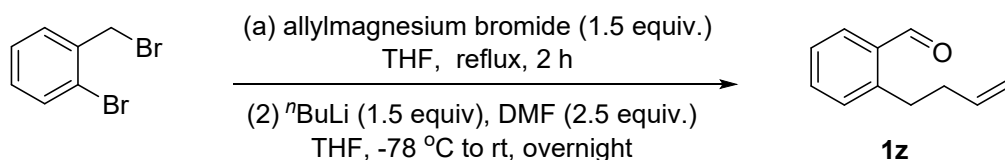


Photoreactor (GeAo)

(B) General procedure for the synthesis of compound 1

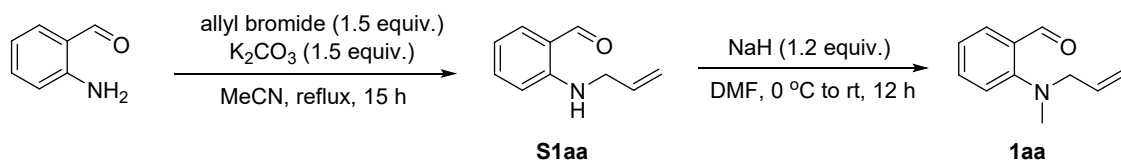


According to the literature.¹ To a solution of 2-hydroxyarylaldehyde (5 mmol, 1.0 equiv.) in DMF (15 mL) was added K₂CO₃ (828 mg, 6 mmol, 1.2 equiv.), followed by the dropwise addition of allyl bromide (726 mg, 6 mmol, 1.2 equiv.). The reaction mixture was then stirred for 12 h at room temperature, poured into brine and extracted with EtOAc. The combined extracts were dried over MgSO₄, filtered, and evaporated. The residue was purified by column chromatography (petroleum ether/ethyl acetate) to afford the desired 2-(allyloxy)arylaldehyde **1a-1y, 1ca**.



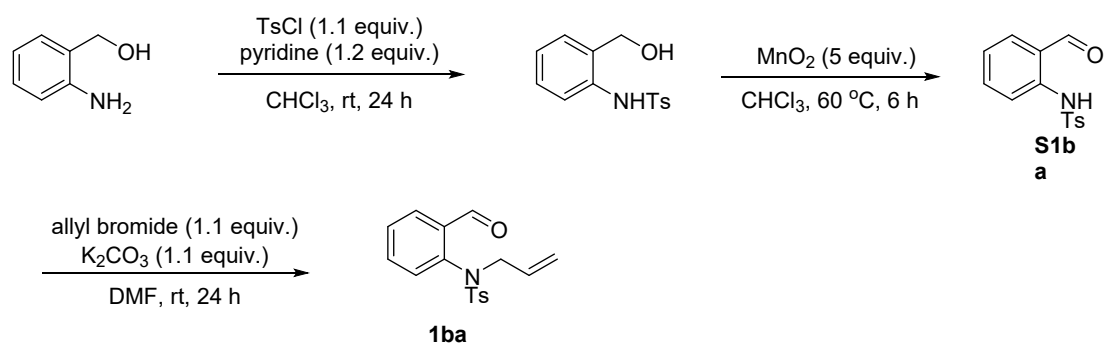
According to the literature.² To a solution of 2-bromobenzyl bromide (2.0 g, 7.3 mmol, 1.0 equiv.) in anhydrous THF (6 mL) was added allylmagnesium bromide (11 mL, 1 M in THF, 11.0 mmol, 1.5 equiv.) at room temperature. The reaction mixture was heated to reflux in an oil bath for 2 h. The reaction mixture was cooled down to room temperature and quenched with saturated NH₄Cl (10 mL), extracted with diethyl ether (3 x 50 mL), dried over anhydrous magnesium sulfate, filtered and concentrated *in vacuo*, the remaining crude product was used directly for the next step without purification.

The crude product was redissolved in anhydrous THF (8 mL) at -78 °C and *n*-BuLi (6.9 mL, 1.6 M in hexanes, 11.0 mmol, 1.5 equiv.) was added dropwise over 30 mins followed by the dropwise addition of a solution of DMF (1.3 mL, 18.2 mmol, 2.5 equiv.) in THF (3.0 mL) over 30 mins. The reaction mixture solution was allowed to warm to room temperature overnight before quenching with saturated NH₄Cl (10 mL). The aqueous layer was extracted with diethyl ether (3 x 50 mL), dried over anhydrous magnesium sulfate, filtered, concentrated *in vacuo* and purified by flash column chromatography (petroleum ether/diethyl ether = 50:1) to afford **1z** (759 mg, 65%) as a colorless oil.



According to the literature.³ To a solution of 2-aminobenzaldehyde (5 mmol, 1.0 equiv.) in anhydrous MeCN (25 mL) was added K₂CO₃ (1.04 g, 7.5 mmol, 1.5 equiv.), followed by the dropwise addition of allyl bromide (907 mg, 7.5 mmol, 1.5 equiv.). The reaction mixture was heated to reflux in an oil bath for 15 h, quenched with saturated NaCl (10 mL) and extracted with EtOAc (3 x 20 mL). The combined extracts were dried over MgSO₄, filtered, and evaporated. The crude product was then purified by silica gel chromatography (petroleum ether/ethyl acetate = 20:1) to afford **S1ca** (363 mg, 61% yield) as a colorless oil.

To a solution of **S1ca** in anhydrous DMF (20 mL) was added NaH (60% dispersion in mineral oil, 164 mg, 4.1 mmol, 1.2 equiv.) was added portionwise at 0 °C. And the reaction mixture was stirred at 0 °C for 10 min. Then MeI (582 mg, 4.1 mmol, 1.2 equiv.) was added to at 0 °C, and then reaction mixture was warmed to room temperature for 12 h. After completion of the reaction, the reaction mixture was quenched with H₂O (5 mL) at 0 °C, extracted with EtOAc (3 x 20 mL). The combined extracts were dried over MgSO₄, filtered, and evaporated. The crude product was then purified by silica gel chromatography (petroleum ether/ethyl acetate = 20:1) to afford **1ca** (578 mg, 72% yield) as a yellow oil.



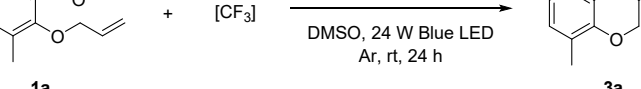
According to the literature.⁴ To a solution of 2-aminobenzyl alcohol (1.23 g, 10 mmol, 1.0 equiv.) in CHCl₃ (35 mL) were added pyridine (949 mg, 12 mmol, 1.2 equiv.) and TsCl (2.10 g, 11 mmol, 1.1 equiv.). The mixture was stirred at room temperature for 24 h to complete the reaction. Then concentrated in vacuo, the remaining crude product was used directly for the next step without purification.

To a solution of the crude product in CHCl₃ (25 mL) was added activated MnO₂ (4.35 g, 50 mmol, 5 equiv.) at room temperature. The reaction mixture was warmed to 60 °C in an oil bath for 6 h. After completed, the mixture was cooled to room temperature and filtered. The filtrate was evaporated under reduced pressure and the crude product was purified by silica gel chromatography (petroleum ether/ethyl acetate = 8:1) to afford **S1aa** (1.68 g, 61% yield) as a white solid.

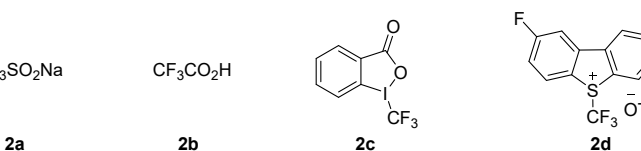
To a solution of **S1aa** in DMF (10 mL) was added K₂CO₃ (881 mg, 6.4 mmol, 1.1 equiv.), followed by the dropwise addition of allyl bromide (772 mg, 6.4 mmol, 1.1 equiv.). The reaction mixture was stirred for 24 h at room temperature. Water (50 mL) was then added and the mixture was extracted with EtOAc (3 × 25 mL). The combined organic extracts were washed with brine (50 mL), dried over anhydrous Na₂SO₄ and concentrated under reduced pressure. The crude product was then purified by silica gel chromatography (petroleum ether/ethyl acetate = 8:1) to afford **1aa** (1.03 g, 56% yield) as a white solid.

(C) Optimization of the reaction conditions

Table S1. Screening of trifluoromethyl reagents^a



Reaction scheme showing the conversion of **1a** to **3a** using $[\text{CF}_3]$ and 4CzIPN (2 mol%) in DMSO, irradiated with 24 W Blue LED, at room temperature for 24 h.



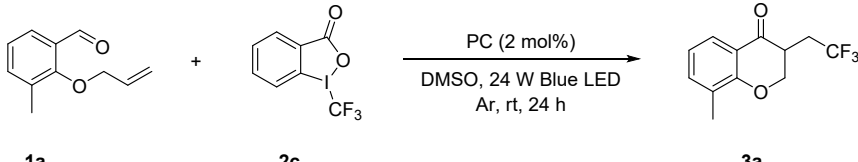
Chemical structures of the reagents used in the reaction:

- 2a**: $\text{CF}_3\text{SO}_2\text{Na}$
- 2b**: $\text{CF}_3\text{CO}_2\text{H}$
- 2c**: 1-(trifluoromethyl)-2,3-dihydro-1H-benzofuran-3-one
- 2d**: 1-(trifluoromethyl)-2,3-dihydro-1H-benzofuran-3-one with a triflate group (OTf)

entry	$[\text{CF}_3]$	yield (%) ^b
1	2a	40
2	2b	n.r.
3	2c	53
4	2d	51

^aReaction conditions: **1a** (0.2 mmol), **2** (0.3 mmol), 4CzIPN (2 mol%), solvent (2.0 mL), irradiation with 24 W blue LED at rt under Ar atmosphere for 24 h. ^bNMR yield were determined by ¹H NMR analysis using 1,3,5-trimethoxybenzene as internal standard.

Table S2. Screening photocatalysts^a

		
1a	2c	3a
entry	PC	yield (%) ^b
1	4CZIPN	53
2	4DPAIPN	50
3	4CZPN-Ph	31
4	3DPAFIPN	49

5	4CZPN-Bu	53
6	Acr ⁺ -MeS ⁺ ClO ₄ ⁻	47
7	[Ir{dFCF ₃ ppy} ₂ (bpy)]PF ₆	34
8	Ru(dmbpy) ₃](PF ₆) ₂	31

^aReaction conditions: **1a** (0.2 mmol), **2c** (0.3 mmol), photocatalyst (2 mol%), DMSO (2.0 mL), irradiation with 24 W blue LED at rt under Ar atmosphere for 24 h. ^bNMR yield were determined by ¹H NMR analysis using 1,3,5-trimethoxybenzene as internal standard.

Table S3. Screening solvents^a

1a	2c	3a	
entry	solvent	yield (%) ^b	
1	DMSO	53	
2	DMF	49	
3	EA	trace	
4	1,4-Dioxane	45	
5	THF	trace	
6	DME	trace	
7	CH ₃ CN	40	
8	CH ₃ OH	36	

^aReaction conditions: **1a** (0.2 mmol), **2c** (0.3 mmol), 4CzIPN (2 mol%), solvent (2.0 mL), irradiation with 24 W blue LED at rt under Ar atmosphere for 24 h. ^bNMR yield were determined by ¹H NMR analysis using 1,3,5-trimethoxybenzene as internal standard.

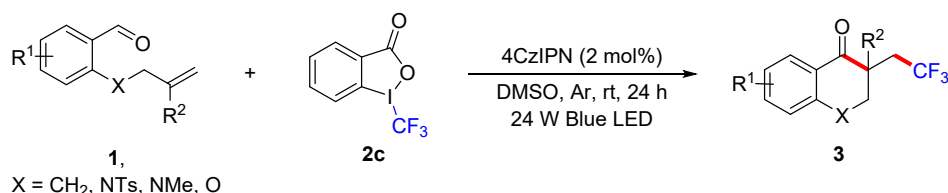
Table S4. Screening the ratio of **1a/2c**^a

1a	2c	3a	
entry	1a (x mmol)	2c (y mmol)	yield (%) ^b
1	1	1	69
2	1	1.5	53
3	1	2	37

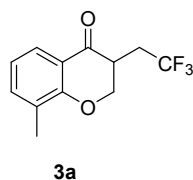
4	1.5	1	81 (78) ^c
5	2	1	83

^aReaction conditions: **1a** (x mmol), **2c** (y mmol), 4CzIPN (2 mol%), DMSO (2.0 mL), irradiation with 24 W blue LED at rt under Ar atmosphere for 24 h. ^bNMR yield were determined by ¹H NMR analysis using 1,3,5-trimethoxybenzene as internal standard. ^cIsolated yield

(D) General procedure for the synthesis of **3**

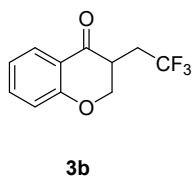


To a screw-capped vial with stirring bar were added 4CzIPN (3.2 mg, 0.004 mmol, 2 mol%), **1** (0.3 mmol, 1.5 equiv.), Togni II reagent **2c** (0.2 mmol, 1.0 equiv.) in an argon-filled glovebox. After the DMSO (2.0 mL) was added into the vial *via* a syringe, the vial was sealed with a cap containing a PTFE septum and removed out from the glovebox. The reaction mixture was stirred at room temperature for 24 h upon irradiation with 24 W blue LED under argon atmosphere. After completion of the reaction, quenched with saturated NaCl (2 mL) and extracted with EtOAc (3 x 5 mL). The combined extracts were dried over MgSO₄, filtered, and evaporated. The crude product was then purified by silica gel chromatography (petroleum ether/ethyl acetate) to afford **3aa-3ca**.



(S)-8-methyl-3-(2,2,2-trifluoroethyl)chroman-4-one (**3a**)

38.1 mg, 78% yield. White solid, mp 67–68 °C, R_f = 0.3 (petroleum ether/ethyl acetate = 80:1). ¹H NMR (400 MHz, CDCl₃) δ 7.75 (dd, *J* = 8.0, 1.2 Hz, 1H), 7.36 (dd, *J* = 7.2, 0.8 Hz, 1H), 6.96–6.92 (m, 1H), 4.77 (dd, *J* = 11.2, 5.2 Hz, 1H), 4.23 (t, *J* = 11.9 Hz, 1H), 3.17–3.01 (m, 2H), 2.25 (s, 3H), 2.15–2.01 (m, 1H). ¹³C NMR (100 MHz, CDCl₃) δ 191.1, 159.8, 137.2, 127.6 (q, *J* = 274.7 Hz), 127.2, 125.2, 121.2, 119.7, 69.7 (d, *J* = 2.5 Hz), 40.4 (d, *J* = 1.7 Hz), 29.6 (q, *J* = 29.9 Hz), 15.5. ¹⁹F NMR (376 MHz, CDCl₃) δ -64.0 (s). HRMS (ESI) *m/z*: [M+H]⁺ calcd for C₁₂H₁₂F₃O₂: 245.0784; Found: 245.0794.

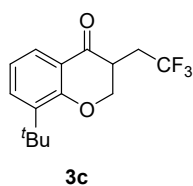


3-(2,2,2-trifluoroethyl)chroman-4-one (**3b**).

25.6 mg, 57% yield. White solid, mp 48–49 °C, R_f = 0.3 (petroleum ether/ethyl acetate = 120:1). ¹H NMR (400 MHz, CDCl₃) δ 7.90 (dd, *J* = 8.0, 1.6 Hz, 1H), 7.53–7.49 (m, 1H), 7.10–7.03 (m, 1H), 6.99 (d, *J* = 8.0 Hz), 4.72 (dd, *J* = 11.4, 5.0 Hz, 1H), 4.24 (t, *J* = 11.8 Hz, 1H), 3.20–3.02 (m, 2H), 2.16–2.01 (m, 1H). ¹³C NMR (100 MHz, CDCl₃) δ 190.9, 161.7, 136.5, 127.7, 126.8 (q, *J* = 274.7 Hz), 121.9, 120.2, 118.0, 69.9 (d,

$J = 1.5$ Hz), 40.6 (d, $J = 2.0$ Hz), 29.7 (q, $J = 30.0$ Hz). **^{19}F NMR (376 MHz, CDCl_3)** δ -64.0 (s).

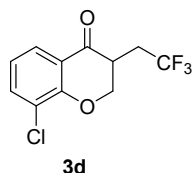
HRMS (ESI) m/z : $[\text{M}+\text{Na}]^+$ calcd for $\text{C}_{11}\text{H}_9\text{F}_3\text{O}_2\text{Na}$: 253.0447; Found: 253.0441.



8-(*tert*-butyl)-3-(2,2,2-trifluoroethyl)chroman-4-one (3c).

36.0 mg, 62% yield. Colorless oil, $R_f = 0.4$ (petroleum ether/ethyl acetate = 40:1). **^1H NMR (400 MHz, CDCl_3)** δ 7.81 (dd, $J = 8.0, 1.6$ Hz, 1H), 7.50 (dd, $J = 7.6, 1.6$ Hz, 1H), 7.00–6.96 (m), 4.79 (dd, $J = 11.0, 5.0$ Hz, 1H), 4.22 (t, $J =$

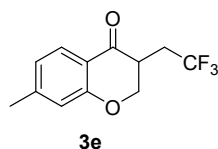
11.6 Hz, 1H), 3.19–3.02 (m, 2H), 2.16–2.02 (m), 1.39 (s, 9H). **^{13}C NMR (100 MHz, CDCl_3)** δ 191.4, 160.8, 139.1, 133.4, 126.7 (q, $J = 274.7$ Hz), 125.7, 121.4, 120.9, 69.3, 40.4 (d, $J = 1.9$ Hz), 34.9, 29.8 (q, $J = 29.8$ Hz). **^{19}F NMR (376 MHz, CDCl_3)** δ -64.0 (s). **HRMS (ESI)** m/z : $[\text{M}+\text{Na}]^+$ calcd for $\text{C}_{15}\text{H}_{17}\text{F}_3\text{O}_2\text{Na}$: 309.1073; Found: 309.1073.



Chloro-3-(2,2,2-trifluoroethyl)chroman-4-one (3d).

29.2 mg, 55% yield. White solid, mp 87–88 °C, $R_f = 0.4$ (petroleum ether/ethyl acetate = 40:1). **^1H NMR (400 MHz, CDCl_3)** δ 7.83 (dd, $J = 8.0, 1.2$ Hz, 1H), 7.60 (dd, $J = 7.8, 1.4$ Hz, 1H), 7.03–6.99 (m, 1H), 4.88 (dd, $J = 11.6, 5.2$ Hz,

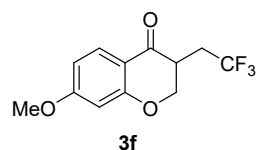
1H), 4.32 (t, $J = 12.0$ Hz, 1H), 3.24–3.02 (m, 2H), 2.17–2.03 (m, 1H). **^{13}C NMR (100 MHz, CDCl_3)** δ 190.1, 157.2, 136.6, 126.6 (q, $J = 274.7$ Hz), 126.3, 122.9, 122.1, 121.5, 70.4, 40.4 (d, $J = 1.9$ Hz), 29.6 (q, $J = 30.1$ Hz). **^{19}F NMR (376 MHz, CDCl_3)** δ -64.0 (s). **HRMS (ESI)** m/z : $[\text{M}+\text{Na}]^+$ calcd for $\text{C}_{11}\text{H}_8\text{ClF}_3\text{O}_2\text{Na}$: 287.0057; Found: 257.0065.



7-methyl-3-(2,2,2-trifluoroethyl)chroman-4-one (3e).

29.3 mg, 60% yield. White solid, mp 62–63 °C, $R_f = 0.3$ (petroleum ether/ethyl acetate = 80:1). **^1H NMR (400 MHz, CDCl_3)** δ 7.78 (d, $J = 8.0$

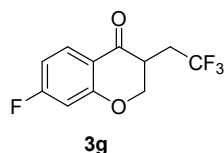
Hz, 1H), 6.86 (d, $J = 8.4$ Hz, 1H), 6.79 (s, 1H), 4.68 (dd, $J = 11.8, 5.0$ Hz, 1H), 4.21 (t, $J = 11.6$ Hz, 1H), 3.14–3.00 (m, 2H), 2.36 (s, 3H), 2.13–1.99 (m, 1H). **^{13}C NMR (100 MHz, CDCl_3)** δ 190.5, 161.8, 148.2, 127.6, 126.8 (q, $J = 274.7$ Hz), 123.4, 118.0, 69.9 (d, $J = 1.4$ Hz), 40.6 (d, $J = 1.9$ Hz), 29.8 (q, $J = 29.9$ Hz), 22.1. **^{19}F NMR (376 MHz, CDCl_3)** δ -64.0 (s). **HRMS (ESI)** m/z : $[\text{M}+\text{H}]^+$ calcd for $\text{C}_{12}\text{H}_{12}\text{F}_3\text{O}_2$: 245.0784; Found: 245.0777.



7-methoxy-3-(2,2,2-trifluoroethyl)chroman-4-one (3f).

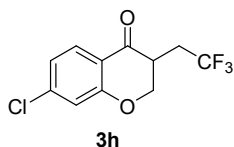
26.7 mg, 48% yield. White solid, mp 100–101 °C, $R_f = 0.4$ (petroleum ether/ethyl acetate = 60:1). **^1H NMR (400 MHz, CDCl_3)** δ 7.83 (d, $J = 8.8$ Hz, 1H), 6.60 (dd, $J = 8.8, 2.4$ Hz, 1H), 6.41 (d, $J = 2.4$ Hz, 1H), 4.69

(dd, $J = 11.2, 4.8$ Hz, 1H), 4.22 (t, $J = 11.4$ Hz, 1H), 3.84 (s, 3H), 3.14–3.01 (m, 2H), 2.12–2.01 (m, 1H). **^{13}C NMR (100 MHz, CDCl_3)** δ 189.4, 166.4, 163.7, 129.5, 126.9 (q, $J = 277.5$ Hz), 114.0, 110.6, 100.8, 70.2, 55.8, 40.2, 29.8 (q, $J = 30$ Hz). **^{19}F NMR (376 MHz, CDCl_3)** δ -64.0 (s). **HRMS (ESI)** m/z : $[\text{M}+\text{H}]^+$ calcd for $\text{C}_{12}\text{H}_{12}\text{F}_3\text{O}_3$: 261.0733; Found: 261.0769.



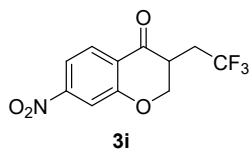
7-fluoro-3-(2,2,2-trifluoroethyl)chroman-4-one (3g).

24.2 mg, 49% yield. White solid, mp 43–44 °C, $R_f = 0.3$ (petroleum ether/ethyl acetate = 120:1). **^1H NMR (400 MHz, CDCl_3)** δ 7.93 (dd, $J = 8.8, 6.8$ Hz, 1H), 6.77 (td, $J = 8.6, 2.4$ Hz, 1H), 6.68 (dd, $J = 9.6, 2.4$ Hz, 1H), 4.74 (dd, $J = 11.4, 5.0$ Hz, 1H), 4.25 (t, $J = 11.8$ Hz, 1H), 3.18–3.02 (m, 2H), 2.14–2.00 (m, 1H). **^{13}C NMR (100 MHz, CDCl_3)** δ 189.5, 167.8 (dd, $J = 255.6$ Hz), 163.4 (d, $J = 13.6$ Hz), 130.4 (d, $J = 11.5$ Hz), 126.7 (q, $J = 274.6$ Hz), 117.1 (d, $J = 2.4$ Hz), 110.5 (d, $J = 22.8$ Hz), 104.9 (d, $J = 22.6$ Hz), 70.3, 40.4 (d, $J = 2.0$ Hz), 29.6 (q, $J = 29.9$ Hz). **^{19}F NMR (376 MHz, CDCl_3)** δ -64.0 (s), -99.4 (s). **HRMS (ESI)** m/z : $[\text{M}+\text{H}]^+$ calcd for $\text{C}_{11}\text{H}_9\text{F}_4\text{O}_2$: 249.0533; Found: 249.0535.



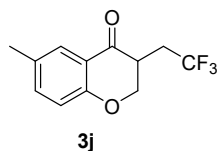
7-chloro-3-(2,2,2-trifluoroethyl)chroman-4-one (3h).

27.6 mg, 52% yield. White solid, mp 63–64 °C, $R_f = 0.4$ (petroleum ether/ethyl acetate = 40:1). **^1H NMR (400 MHz, CDCl_3)** δ 7.85–7.83 (m, 1H), 7.04 (dd, $J = 6.6, 1.8$ Hz, 2H), 4.74 (dd, $J = 11.2, 5.2$ Hz, 1H), 4.24 (t, $J = 11.8$ Hz, 1H), 3.18–3.00 (m, 2H), 2.15–2.01 (m, 1H). **^{13}C NMR (100 MHz, CHCl_3)** δ 189.8, 162.0, 142.5, 129.0, 126.7 (q, $J = 274.8$ Hz), 122.8, 118.8, 118.2, 77.2, 76.8, 29.7 (q, $J = 30.1$ Hz). **^{19}F NMR (376 MHz, CDCl_3)** δ -64.0 (s). **HRMS (ESI)** m/z : $[\text{M}+\text{H}]^+$ calcd for $\text{C}_{11}\text{H}_9\text{ClF}_3\text{O}_2$: 265.0238; Found: 265.0238.



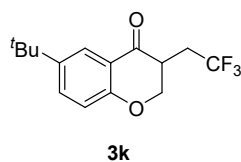
7-nitro-3-(2,2,2-trifluoroethyl)chroman-4-one (3i).

18.2 mg, 33% yield. Orange oil, $R_f = 0.4$ (petroleum ether/ethyl acetate = 20:1). **^1H NMR (400 MHz, CDCl_3)** δ 8.09–8.06 (m, 1H), 7.86 (dd, $J = 6.8, 2.0$ Hz, 2H), 4.85 (dd, $J = 11.6, 5.2$ Hz, 1H), 4.33 (t, $J = 12.0$ Hz, 1H), 3.29–3.22 (m, 1H), 3.18–3.04 (m, 1H), 2.19–2.04 (m, 1H). **^{13}C NMR (100 MHz, CDCl_3)** δ 189.6, 161.7, 152.5, 129.4, 126.4 (q, $J = 274.7$ Hz), 123.9, 116.3, 114.0, 70.3, 40.7 (d, $J = 2.0$ Hz), 29.5 (q, $J = 30.3$ Hz). **^{19}F NMR (376 MHz, CDCl_3)** δ -64.0 (s). **HRMS (ESI)** m/z : $[\text{M}+\text{Na}]^+$ calcd for $\text{C}_{11}\text{H}_8\text{F}_3\text{NO}_4\text{Na}$: 298.0298; Found: 298.0286.



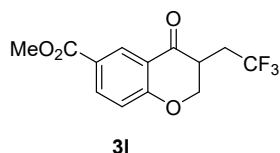
6-methyl-3-(2,2,2-trifluoroethyl)chroman-4-one (3j).

28.4 mg, 58% yield. White solid, mp 68–69 °C, R_f = 0.3 (petroleum ether/ethyl acetate = 120:1). $^1\text{H NMR}$ (400 MHz, CDCl_3) δ 7.69 (d, J = 1.2 Hz, 1H), 7.32 (dd, J = 8.4, 2.0 Hz, 1H), 6.89 (d, J = 8.4 Hz, 1H), 4.68 (dd, J = 11.0, 5.0 Hz, 1H), 4.21 (t, J = 11.6 Hz, 1H), 3.16–3.00 (m, 2H), 2.32 (s, 3H), 2.15–2.01 (m, 1H). $^{13}\text{C NMR}$ (100 MHz, CDCl_3) δ 191.0, 159.7, 137.5, 131.3, 127.1, 126.6 (q, J = 274.6 Hz), 119.7, 117.7, 69.8, 40.6, 29.7 (q, J = 29.9 Hz), 20.4. $^{19}\text{F NMR}$ (376 MHz, CDCl_3) δ -64.0 (s). HRMS (ESI) m/z : $[\text{M}+\text{H}]^+$ calcd for $\text{C}_{12}\text{H}_{12}\text{F}_3\text{O}_2$: 245.0784; Found: 245.0786.



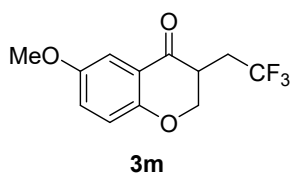
6-(tert-butyl)-3-(2,2,2-trifluoroethyl)chroman-4-one (3k).

31.5 mg, 55% yield, White solid, mp 62–63 °C, R_f = 0.4 (petroleum ether/ethyl acetate = 40:1). $^1\text{H NMR}$ (400 MHz, CDCl_3) δ 7.89 (d, J = 2.4 Hz, 1H), 7.57 (dd, J = 8.6, 2.6 Hz, 1H), 6.93 (d, J = 8.8 Hz, 1H), 4.68 (dd, J = 11.4, 5.0 Hz, 1H), 4.22 (m, J = 11.8 Hz, 1H), 3.16–3.00 (m, 2H), 2.15–2.01 (m, 1H), 1.31 (s, 9H). $^{13}\text{C NMR}$ (100 MHz, CDCl_3) δ 191.2, 159.7, 144.9, 134.3, 126.8 (q, J = 274.7 Hz), 123.6, 119.4, 117.6, 69.9 (d, J = 1.3 Hz), 40.7 (d, J = 1.9 Hz), 34.5, 31.4, 29.8 (q, J = 29.9 Hz). $^{19}\text{F NMR}$ (376 MHz, CDCl_3) δ -64.0 (s). HRMS (ESI) m/z : $[\text{M}+\text{Na}]^+$ calcd for $\text{C}_{15}\text{H}_{17}\text{F}_3\text{O}_2\text{Na}$: 309.1073; Found: 309.1082.



methyl 4-oxo-3-(2,2,2-trifluoroethyl)chromane-6-carboxylate (3l).

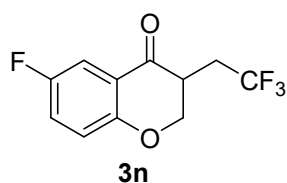
36.4 mg, 63% yield, White solid, mp 136–137 °C, R_f = 0.3 (petroleum ether/ethyl acetate = 20:1). $^1\text{H NMR}$ (400 MHz, CDCl_3) δ 8.57 (d, J = 2.4 Hz, 1H), 8.15 (dd, J = 8.8, 2.4 Hz, 1H), 7.03 (d, J = 8.4 Hz, 1H), 4.78 (dd, J = 11.6, 5.2 Hz, 1H), 4.28 (t, J = 11.8 Hz, 1H), 3.90 (s, 3H), 3.22–3.03 (m, 2H), 2.16–2.02 (m, 1H). $^{13}\text{C NMR}$ (100 MHz, CDCl_3) δ 184.5, 160.6, 159.3, 131.8, 124.8, 121.3 (q, J = 274.7 Hz), 118.9, 114.4, 113.0, 64.7, 47.0, 35.2 (d, J = 1.9 Hz), 24.3 (q, J = 30.1 Hz). $^{19}\text{F NMR}$ (376 MHz, CDCl_3) δ -64.0 (s). HRMS (ESI) m/z : $[\text{M}+\text{Na}]^+$ calcd for $\text{C}_{13}\text{H}_{11}\text{F}_3\text{O}_4\text{Na}$: 311.0502; Found: 311.0501.



(S)-6-methoxy-3-(2,2,2-trifluoroethyl)chroman-4-one (3m)

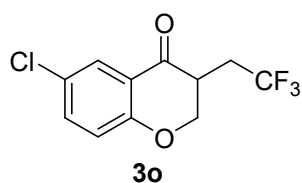
24.9 mg, 48% yield. Colorless oil, R_f = 0.3 (petroleum ether/ethyl acetate = 20:1). $^1\text{H NMR}$ (400 MHz, CDCl_3) δ 7.30 (d, J = 3.2 Hz, 1H), 7.11 (dd, J = 8.8, 3.2 Hz, 1H), 6.92 (d, J = 8.8 Hz, 1H), 4.67 (dd, J = 11.6, 5.2 Hz, 1H), 4.20 (t, J = 11.6 Hz, 1H), 3.80 (s, 3H), 3.16–2.99 (m, 2H), 2.16–2.01 (m,

1H). ¹³C NMR (100 MHz, CDCl₃) δ 191.0, 156.4, 154.5, 125.8, 125.7 (q, *J* = 274.8 Hz), 120.0, 119.3, 107.9, 70.0 (d, *J* = 1.3 Hz), 55.9, 40.6 (d, *J* = 1.9 Hz), 29.8 (q, *J* = 29.9 Hz). ¹⁹F NMR (376 MHz, CDCl₃) δ -64.0 (s). HRMS (ESI) *m/z*: [M+H]⁺ calcd for C₁₂H₁₂F₃O₃: 261.0733; Found: 261.0739.



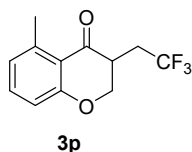
(S)-6-fluoro-3-(2,2,2-trifluoroethyl)chroman-4-one (3n)

34.3 mg, 69% yield. White solid, mp 74–75 °C, *R*_f = 0.3 (petroleum ether/ethyl acetate = 120:1). ¹H NMR (400 MHz, CDCl₃) δ 7.56 (dd, *J* = 8.4, 3.2 Hz, 1H), 7.28–7.23 (m, 1H), 7.00 (dd, *J* = 9.0, 4.2 Hz, 1H), 4.73 (dd, *J* = 11.2, 5.2 Hz, 1H), 4.24 (t, *J* = 11.8 Hz, 1H), 3.20–3.02 (m, 2H), 2.18–2.03 (m, 1H). ¹³C NMR (100 MHz, CDCl₃) δ 190.2, 158.4 (d, *J* = 75.4 Hz), 156.3, 126.6 (q, *J* = 274.7 Hz), 124.2 (d, *J* = 24.4 Hz), 120.6 (d, *J* = 6.6 Hz), 119.8 (d, *J* = 7.4 Hz), 112.6 (d, *J* = 23.3 Hz), 70.1 (d, *J* = 1.4 Hz), 40.6, 29.7 (q, *J* = 30.0 Hz). ¹⁹F NMR (376 MHz, CDCl₃) δ -64.0 (s), -120.7 (s). HRMS (ESI) *m/z*: [M+H]⁺ calcd for C₁₁H₉F₄O₂: 249.0533; Found: 249.0532.



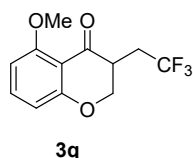
(S)-6-chloro-3-(2,2,2-trifluoroethyl)chroman-4-one (3o)

40.3 mg, 76% yield. White solid, mp 70–71 °C, *R*_f = 0.3 (petroleum ether/ethyl acetate = 100:1). ¹H NMR (400 MHz, CDCl₃) δ 7.86 (d, *J* = 2.8 Hz, 1H), 7.44 (dd, *J* = 8.8, 2.8 Hz, 1H), 6.96 (d, *J* = 8.8 Hz, 1H), 4.74 (dd, *J* = 11.6, 5.2 Hz, 1H), 4.23 (t, *J* = 11.8 Hz, 1H), 3.18–3.00 (m, 2H), 2.16–2.01 (m, 1H). ¹³C NMR (100 MHz, CDCl₃) δ 189.8, 160.2, 136.4, 127.6, 127.0, 126.6 (q, *J* = 273.5 Hz), 121.0, 119.8, 70.0 (d, *J* = 1.4 Hz), 40.5 (d, *J* = 1.9 Hz), 29.7 (q, *J* = 30.1 Hz). ¹⁹F NMR (376 MHz, CDCl₃) δ -64.0 (s). HRMS (ESI) *m/z*: [M+Na]⁺ calcd for C₁₁H₈ClF₃O₂Na: 287.0057; Found: 257.0067.



5-methyl-3-(2,2,2-trifluoroethyl)chroman-4-one (3p).

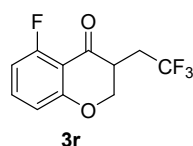
26.5 mg, 54% yield. White solid, mp 54–55 °C, *R*_f = 0.3 (petroleum ether/ethyl acetate = 120:1). ¹H NMR (400 MHz, CDCl₃) δ 7.35–7.31 (m, 1H), 6.85–6.81 (m, 2H), 4.66 (dd, *J* = 11.2, 4.8 Hz, 1H), 4.20 (t, *J* = 11.6 Hz, 1H), 3.15–2.97 (m, 2H), 2.62 (s, 3H), 2.14–2.00 (m, 1H). ¹³C NMR (100 MHz, CDCl₃) δ 191.1, 161.6, 141.6, 134.2, 125.9 (q, *J* = 274.7 Hz), 124.1, 117.8, 114.9, 68.2 (d, *J* = 1.6 Hz), 40.4 (d, *J* = 1.7 Hz), 28.8 (q, *J* = 29.7 Hz), 22.0. ¹⁹F NMR (376 MHz, CDCl₃) δ -64.0 (s). HRMS (ESI) *m/z*: [M+H]⁺ calcd for C₁₂H₁₂F₃O₂: 245.0784; Found: 245.0792.



5-methoxy-3-(2,2,2-trifluoroethyl)chroman-4-one (3q).

32.5 mg, 63% yield. Yellow oil, R_f = 0.3 (petroleum ether/ethyl acetate = 20:1).

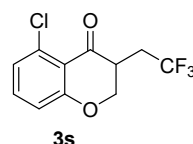
^1H NMR (400 MHz, CDCl_3) δ 7.41–7.37 (m, 1H), 6.58–6.52 (m, 2H), 4.64 (dd, J = 11.6, 4.8 Hz, 1H), 4.19 (t, J = 11.2 Hz, 1H), 3.90 (s, 3H), 3.12–2.98 (m, 2H), 2.12–1.97 (m, 1H). ^{13}C NMR (100 MHz, CDCl_3) δ 189.1, 163.0, 161.0, 136.3, 126.8 (q, J = 274.6 Hz), 110.5, 109.8, 104.1, 69.2 (d, J = 1.3 Hz), 56.2, 41.3 (d, J = 2.0 Hz), 29.6 (q, J = 29.6 Hz). ^{19}F NMR (376 MHz, CDCl_3) δ -64.0 (s). HRMS (ESI) m/z : $[\text{M}+\text{H}]^+$ calcd for $\text{C}_{12}\text{H}_{12}\text{F}_3\text{O}_3$: 261.0733; Found: 261.0725.



5-fluoro-3-(2,2,2-trifluoroethyl)chroman-4-one (3r).

38.1 mg, 77% yield. White solid, mp 82–83 °C, R_f = 0.3 (petroleum ether/ethyl acetate = 20:1).

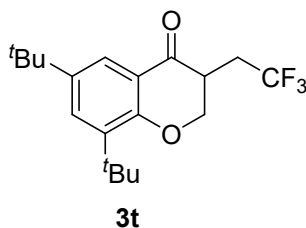
^1H NMR (400 MHz, CDCl_3) δ 7.46–7.41 (m, 1H), 6.80 (d, J = 8.8 Hz, 1H), 6.75–6.70 (m, 1H), 4.71 (dd, J = 11.6, 5.2 Hz, 1H), 4.24 (t, J = 11.8 Hz, 1H), 3.18–3.00 (m, 2H), 2.14–2.00 (m, 1H). ^{13}C NMR (100 MHz, CDCl_3) δ 188.2, 161.8 (d, J = 264.9 Hz), 162.2 (d, J = 2.9 Hz), 136.4 (d, J = 11.7 Hz), 126.6 (q, J = 274.7 Hz), 113.6 (d, J = 4.0 Hz), 110.3 (d, J = 9.3 Hz), 109.2 (d, J = 10.9 Hz), 69.6 (d, J = 1.5 Hz), 41.1 (d, J = 1.8 Hz), 29.4 (q, J = 29.9 Hz). ^{19}F NMR (376 MHz, CDCl_3) δ -64.0 (s), -110.1 (s). HRMS (ESI) m/z : $[\text{M}+\text{H}]^+$ calcd for $\text{C}_{11}\text{H}_9\text{F}_4\text{O}_2$: 249.0533; Found: 249.0530.



5-chloro-3-(2,2,2-trifluoroethyl)chroman-4-one (3s).

44.8 mg, 85% yield. Colorless oil, R_f = 0.3 (petroleum ether/ethyl acetate = 100:1).

^1H NMR (400 MHz, CDCl_3) δ 7.37–7.33 (m, 1H), 7.06 (d, J = 7.6 Hz, 1H), 6.91 (m, J = 8.4 Hz, 1H), 4.70 (dd, J = 11.2, 5.2 Hz, 1H), 4.23 (t, J = 11.8 Hz, 1H), 3.21–2.99 (m, 2H), 2.14–2.00 (m, 1H). ^{13}C NMR (100 MHz, CDCl_3) δ 188.9, 163.0, 135.3, 134.9, 126.7 (q, J = 274.7 Hz), 125.2, 117.6, 117.1, 69.4, 41.2 (d, J = 1.9 Hz), 29.7 (q, J = 30.0 Hz). ^{19}F NMR (376 MHz, CDCl_3) δ -64.0 (s). HRMS (ESI) m/z : $[\text{M}+\text{H}]^+$ calcd for $\text{C}_{11}\text{H}_9\text{ClF}_3\text{O}_2$: 265.0238; Found: 265.0247.

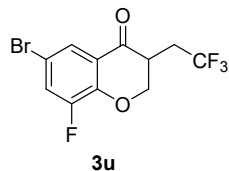


6,8-di-tert-butyl-3-(2,2,2-trifluoroethyl)chroman-4-one (3t).

36.7 mg, 68% yield. White solid, mp 69–70 °C, R_f = 0.3 (petroleum ether/ethyl acetate = 100:1).

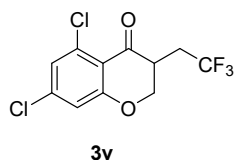
^1H NMR (400 MHz, CDCl_3) δ 7.81 (d, J = 2.4 Hz, 1H), 7.58 (d, J = 2.4 Hz, 1H), 4.75 (dd, J = 11.2, 4.8 Hz, 1H), 4.21 (t, J = 11.6 Hz, 1H), 3.16–3.01 (m, 2H), 2.07–2.02 (m, 5.1 Hz, 1H), 1.40 (s, 9H), 1.32 (s, 9H). ^{13}C NMR (100 MHz, CDCl_3) δ 191.7, 158.7, 143.8, 138.4,

131.1, 126.8 (q, $J = 274.7$ Hz), 121.6, 120.2, 69.3, 40.5 (d, $J = 1.9$ Hz), 35.1, 34.5, 31.3, 29.7, 29.9 (q, $J = 29.8$ Hz). **^{19}F NMR (376 MHz, CDCl_3)** δ -63.9 (s). **HRMS (ESI)** m/z : $[\text{M}+\text{Na}]^+$ calcd for $\text{C}_{19}\text{H}_{25}\text{F}_3\text{O}_2\text{Na}$: 365.1699; Found: 365.1693.



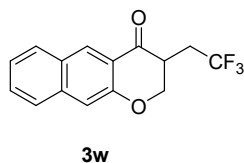
6-bromo-8-fluoro-3-(2,2,2-trifluoroethyl)chroman-4-one (3u).

47.7 mg, 74% yield. White solid, mp 99–100 °C, $R_f = 0.3$ (petroleum ether/ethyl acetate = 120:1). **^1H NMR (400 MHz, CDCl_3)** δ 7.81 (d, $J = 1.6$ Hz, 1H), 7.47–7.44 (m, 1H), 4.84 (dd, $J = 11.6, 5.2$ Hz, 1H), 4.30 (t, $J = 11.8$ Hz, 1H), 3.24–3.00 (m, 2H), 2.18–2.04 (m, 1H). **^{13}C NMR (100 MHz, CDCl_3)** δ 188.6 (d, $J = 3.0$ Hz), 151.6 (d, $J = 253.7$ Hz), 149.3 (d, $J = 11.6$ Hz), 126.5 (q, $J = 274.8$ Hz), 125.6, 125.4 (d, $J = 1.9$ Hz), 122.9, 113.2 (d, $J = 7.4$ Hz), 70.6, 40.7 (d, $J = 1.9$ Hz), 29.6 (q, $J = 30.3$ Hz). **^{19}F NMR (376 MHz, CDCl_3)** δ -64.0 (s), -131.3 (s). **HRMS (ESI)** m/z : $[\text{M}+\text{H}]^+$ calcd for $\text{C}_{11}\text{H}_8^{79}\text{BrF}_4\text{O}_2$: 326.9638; Found: 326.9638; for $\text{C}_{11}\text{H}_8^{81}\text{BrF}_4\text{O}_2$: 328.9618; Found: 328.9618.



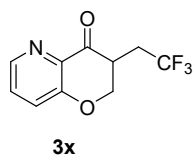
5,7-dichloro-3-(2,2,2-trifluoroethyl)chroman-4-one (3v).

51.7 mg, 87% yield. White solid, mp 84–85 °C, $R_f = 0.3$ (petroleum ether/ethyl acetate = 120:1). **^1H NMR (400 MHz, CDCl_3)** δ 7.08 (d, $J = 1.6$ Hz, 1H), 6.95 (d, $J = 2.0$ Hz, 1H), 4.72 (dd, $J = 11.6, 5.2$ Hz, 1H), 4.24 (t, $J = 11.8$ Hz, 1H), 3.20–2.99 (m, 2H), 3.14–2.00 (m, 1H). **^{13}C NMR (100 MHz, CDCl_3)** δ 187.8, 162.9, 140.9, 135.8, 126.5 (q, $J = 274.7$ Hz), 125.3, 117.1, 116.0, 69.5 (d, $J = 1.4$ Hz), 41.0 (d, $J = 1.8$ Hz), 29.5 (q, $J = 30.0$ Hz). **^{19}F NMR (376 MHz, CDCl_3)** δ -64.0 (s). **HRMS (ESI)** m/z : $[\text{M}+\text{H}]^+$ calcd for $\text{C}_{11}\text{H}_8\text{Cl}_2\text{F}_3\text{O}_2$: 298.9848; Found: 298.9848.



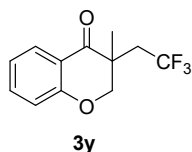
3-(2,2,2-trifluoroethyl)-2,3-dihydro-4H-benzo[g]chromen-4-one (3w).

23.3 mg, 42% yield. White solid, mp 62–63 °C, $R_f = 0.3$ (petroleum ether/ethyl acetate = 120:1). **^1H NMR (400 MHz, CDCl_3)** δ 9.39 (d, $J = 8.4$ Hz, 1H), 7.95 (d, $J = 8.8$ Hz, 1H), 7.77 (d, $J = 8.0$ Hz, 1H), 7.67–7.63 (m, 1H), 7.47–7.44 (m, 1H), 7.11 (d, $J = 8.8$ Hz, 1H), 4.79 (dd, $J = 11.2, 5.2$ Hz, 1H), 4.37 (t, $J = 11.6$ Hz, 1H), 3.27–3.06 (m, 2H), 2.24–2.10 (m, 1H). **^{13}C NMR (100 MHz, CDCl_3)** δ 191.6, 163.7, 138.1, 131.6, 130.0, 129.4, 128.7, 126.9 (q, $J = 274.7$ Hz), 125.8, 125.2, 118.6, 111.9, 69.8, 41.09 (d, $J = 2.0$ Hz), 30.1 (q, $J = 29.6$ Hz). **^{19}F NMR (376 MHz, CDCl_3)** δ -63.8 (s). **HRMS (ESI)** m/z : $[\text{M}+\text{H}]^+$ calcd for $\text{C}_{15}\text{H}_{12}\text{F}_3\text{O}_2$: 281.0784; Found: 281.0786.



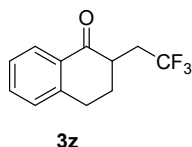
3-(2,2,2-trifluoroethyl)-2,3-dihydro-4H-pyrano[3,2-b]pyridin-4-one (3x).

38.1 mg, 83% yield. Yellow oil, $R_f = 0.4$ (petroleum ether/ethyl acetate = 4:1). **^1H NMR (400 MHz, CDCl_3)** δ 8.50 (dd, $J = 3.8, 1.4$ Hz, 1H), 7.47–7.41 (m, 2H), 4.80 (dd, $J = 11.6, 5.2$ Hz, 1H), 4.32 (t, $J = 12.0$ Hz, 1H), 3.35–3.27 (m, 1H), 3.18 (m, 1H), 3.25–3.12 (m, 1H). **^{13}C NMR (100 MHz, CDCl_3)** δ 189.3, 159.4, 144.8, 136.1, 129.6, 126.8, 126.4 (q, $J = 274.7$ Hz), 69.9 (d, $J = 1.8$ Hz), 41.3 (d, $J = 2.1$ Hz), 29.5 (q, $J = 30.1$ Hz). **^{19}F NMR (376 MHz, CDCl_3)** δ -64.0 (s). **HRMS (ESI)** m/z : $[\text{M}+\text{H}]^+$ calcd for $\text{C}_{10}\text{H}_9\text{F}_3\text{NO}_2$: 232.0580; Found: 232.0580.



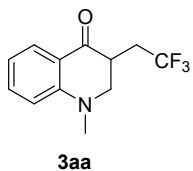
4-methyl-3-(2,2,2-trifluoroethyl)chroman-4-one (3y).

16.2 mg, 34% yield. Colorless oil, $R_f = 0.3$ (petroleum ether/ethyl acetate = 70:1). **^1H NMR (400 MHz, CDCl_3)** δ 7.91 (dd, $J = 7.8, 1.4$ Hz, 1H), 7.53–7.49 (m, 1H), 7.08–7.04 (m, 1H), 6.99 (d, $J = 8.4$ Hz, 1H), 4.35 (q, $J = 11.3$ Hz, 2H), 2.57 (qd, $J = 11.6, 1.2$ Hz, 2H), 1.35 (s, 3H). **^{13}C NMR (100 MHz, CDCl_3)** δ 193.9, 160.8, 136.2, 128.2, 126.3 (q, $J = 276.4$ Hz), 122.0, 118.9, 117.8, 73.9 (d, $J = 2.1$ Hz), 36.3 (q, $J = 28.6$ Hz), 42.6, 18.6. **^{19}F NMR (376 MHz, CDCl_3)** δ -59.5 (s). **HRMS (ESI)** m/z : $[\text{M}+\text{H}]^+$ calcd for $\text{C}_{12}\text{H}_{12}\text{F}_3\text{O}_2$: 245.0784; Found: 245.0775.



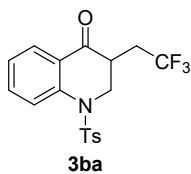
2-(2,2,2-trifluoroethyl)-3,4-dihydronaphthalen-1(2H)-one (3z).

32.3 mg, 71% yield. White solid, mp 42–43 °C, $R_f = 0.3$ (petroleum ether/ethyl acetate = 30:1). **^1H NMR (400 MHz, CDCl_3)** δ 8.04 (d, $J = 7.6$ Hz, 1H), 7.52–7.48 (m, 1H), 7.32 (t, $J = 7.6$ Hz, 1H), 7.27–7.25 (m, 1H), 3.27–3.08 (m, 2H), 3.04–2.97 (m, 1H), 2.86–2.79 (m, 1H), 2.52–2.45 (m, 1H), 2.18–2.03 (m, 1H), 2.01–1.90 (m, 1H). **^{13}C NMR (100 MHz, CDCl_3)** δ 196.7, 143.8, 133.8, 131.9, 128.9, 127.9, 127.4 (q, $J = 274.8$ Hz), 126.9, 42.8 (d, $J = 2.0$ Hz), 33.4 (q, $J = 28.5$ Hz), 29.8, 29.1 (d, $J = 10.8$ Hz). **^{19}F NMR (376 MHz, CDCl_3)** δ -63.6 (s). **HRMS (ESI)** m/z : $[\text{M}+\text{H}]^+$ calcd for $\text{C}_{12}\text{H}_{12}\text{F}_3\text{O}$: 229.0835; Found: 229.0840.



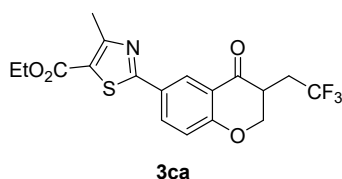
1-methyl-3-(2,2,2-trifluoroethyl)-2,3-dihydroquinolin-4(1H)-one (3aa).

29.9 mg, 62% yield. Yellow solid, mp 59–60 °C, $R_f = 0.3$ (petroleum ether/ethyl acetate = 120:1). **^1H NMR (400 MHz, CDCl_3)** δ 7.91 (dd, $J = 8.0, 1.6$ Hz, 1H), 7.45–7.41 (m, 1H), 6.80–6.72 (m, 2H), 3.61 (dd, $J = 12.0, 5.2$ Hz, 1H), 3.33 (t, $J = 12.4$ Hz, 1H), 3.15–2.97 (m, 5H), 2.14–2.00 (m, 1H). **^{13}C NMR (100 MHz, CDCl_3)** δ 192.2, 152.3, 135.8, 128.6, 127.1 (q, $J = 274.7$ Hz), 118.8, 117.5, 113.2, 55.5, 41.0 (d, $J = 1.7$ Hz), 39.4, 30.8 (q, $J = 29.3$ Hz). **^{19}F NMR (376 MHz, CDCl_3)** δ 63.8 (s). **HRMS (ESI)** m/z : $[\text{M}+\text{H}]^+$ calcd for $\text{C}_{12}\text{H}_{13}\text{F}_3\text{NO}$: 244.0944; Found: 244.0945.



1-tosyl-3-(2,2,2-trifluoroethyl)-2,3-dihydroquinolin-4(1H)-one (3ba).

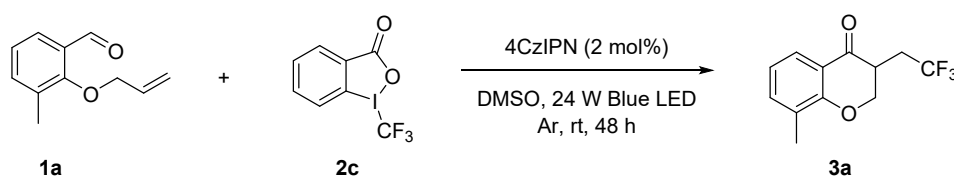
41.3 mg, 54% yield. White solid, mp 103–104 °C, R_f = 0.5 (petroleum ether/ethyl acetate = 10:1). ^1H NMR (400 MHz, CDCl_3) δ 7.90–7.86 (m, 2H), 7.55–7.49 (m, 3H), 7.22–7.49 (m, 3H), 4.68 (dd, J = 14.4, 4.8 Hz, 1H), 3.61 (t, J = 13.9 Hz, 1H), 2.98–2.84 (m, 1H), 2.44–2.37 (m, 1H), 2.32 (s, 3H), 1.89–1.75 (m, 1H). ^{13}C NMR (100 MHz, CDCl_3) δ 191.6, 144.9, 142.2, 136.6, 135.2, 130.3, 128.3, 127.0 (J = 274.9 Hz), 126.8, 125.6, 124.3, 124.1, 50.1, 39.6 (d, J = 1.9 Hz), 30.8 (q, J = 29.7 Hz), 21.6. ^{19}F NMR (376 MHz, CDCl_3) δ 63.8 (s). HRMS (ESI) m/z : $[\text{M}+\text{Na}]^+$ calcd for $\text{C}_{18}\text{H}_{16}\text{F}_3\text{NO}_3\text{SNa}$: 406.0695; Found: 406.0702.



Ethyl 4-methyl-2-(4-oxo-3-(2,2,2-trifluoroethyl)chroman-6-yl)thiazole-5-carboxylate (3ca).

42.1 mg, 58% yield. White solid, mp 127–128 °C, R_f = 0.3 (petroleum ether/ethyl acetate = 8:1). ^1H NMR (400 MHz, CDCl_3) δ 8.41 (d, J = 2.0 Hz, 1H), 8.17 (dd, J = 8.6, 2.2 Hz, 1H), 7.07 (d, J = 8.8 Hz, 1H), 4.78 (dd, J = 11.6, 5.2 Hz, 1H), 4.37–4.32 (m, 2H), 4.32–4.26 (m, 1H), 3.23–3.04 (m, 2H), 2.75 (s, 3H), 2.18–2.04 (m, 1H), 1.38 (t, J = 7.0 Hz, 3H). ^{13}C NMR (100 MHz, CDCl_3) δ 188.8, 167.1, 162.2, 160.5 (d, J = 115.7 Hz), 133.3, 126.1, 125.5 (q, J = 275.0 Hz), 125.3, 120.9, 119.1, 117.9, 68.9, 60.3, 39.4, 30.4, 28.6 (q, J = 30.1 Hz), 16.4, 13.3. ^{19}F NMR (376 MHz, CDCl_3) δ -64.0 (s). HRMS (ESI) m/z : $[\text{M}+\text{H}]^+$ calcd for $\text{C}_{18}\text{H}_{17}\text{F}_3\text{NO}_4\text{S}$: 400.0825; Found: 400.0835.

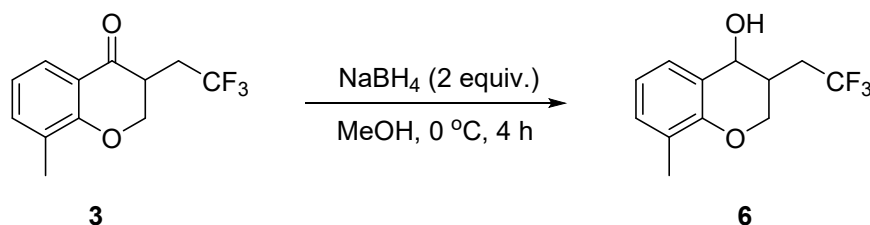
(E) Scale-up reaction and synthetic applications



To a screw-capped vial with stirring bar were added 4CzIPN (3.2 mg, 0.004 mmol, 2 mol%), **1a** (528 mg, 3 mmol, 1.5 equiv.), Togni II reagent **2c** (632 mg, 2 mmol, 1.0 equiv.) in an argon-filled glovebox. After the DMSO (20 mL) was added into the vial *via* a syringe, the vial was sealed with a cap containing a PTFE septum and removed out from the glovebox. The reaction mixture was stirred at room temperature for 48 h upon irradiation with 24 W blue LED under argon atmosphere. After completion of the reaction, quenched with saturated NaCl (20 mL) and extracted with EtOAc (3 x 5 mL). The combined extracts were dried over MgSO_4 , filtered, and evaporated. The crude product was then purified by silica gel chromatography ((petroleum

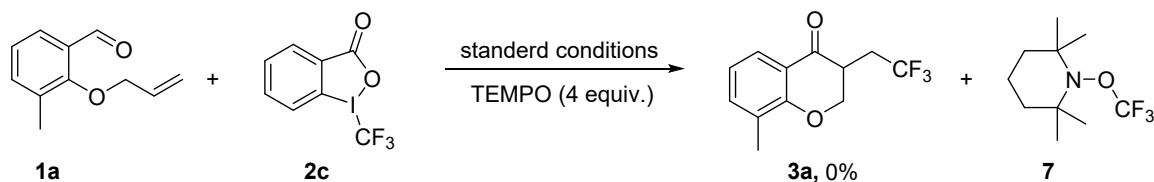
16

afford **34** (35.8 mg, 74% yield) as white solid. **¹H NMR (400 MHz, CDCl₃)** δ 8.08 (d, *J* = 8.0 Hz, 1H), 8.04 (s, 1H), 7.53 (d, *J* = 6.8 Hz, 1H), 7.32 (m, 1H), 3.40 (q, *J* = 10.8 Hz, 2H), 2.48 (s, 3H). **¹³C NMR (100 MHz, CDCl₃)** δ 176.8, 155.0, 154.9, 135.1, 127.8, 125.8 (q, *J* = 275.1 Hz), 125.3, 123.9, 123.6, 114.6, 29.2 (q, *J* = 31.4 Hz), 15.6. **¹⁹F NMR (376 MHz, CDCl₃)** δ -65.8 (s). **HRMS (ESI)** *m/z*: [M+H]⁺ calcd for C₁₂H₁₀F₃O₂: 243.0627; Found: 243.0621.



To a solution of compound **3** (48.8 mg, 0.2 mmol, 1.0 equiv.) in anhydrous MeOH (2 mL) was added NaBH₄ (3.5 mg, 0.4 mmol, 2 equiv.) at 0 °C. And the reaction mixture was stirred at the same temperature for 4 h. Quenched with saturated NaCl (5 mL) and the aqueous layer was extracted with EtOAc (5 mL x 3). The combined organic layers were dried over anhydrous Na₂SO₄, filtered, and concentrated under reduced pressure. The resulting residue was purified by column chromatography (petroleum ether/ethyl acetate = 20 : 1,) to afford **6** (33.3 mg, 68% yield) as a white solid. **¹H NMR (400 MHz, CDCl₃)** δ 7.15 (d, *J* = 7.6 Hz, 1H), 7.11 (d, *J* = 7.6 Hz, 1H), 6.89–6.85 (m, 1H), 4.51 (s, 1H), 4.41 (dd, *J* = 11.2, 1.2 Hz, 1H), 4.17 (dd, *J* = 11.2, 4.0 Hz, 1H), 2.36–2.30 (m, 1H), 2.20 (s, 3H), 2.18–2.06 (m, 2H), 1.93 (s, 1H). **¹³C NMR (100 MHz, CDCl₃)** δ 152.2, 131.3, 127.6, 126.9 (q, *J* = 274.5 Hz), 126.5, 121.7, 120.7, 67.8, 64.7, 34.2, 32.5 (q, *J* = 28.7 Hz), 16.0. **¹⁹F NMR (376 MHz, CDCl₃)** δ -64.0 (s). **HRMS (ESI)** *m/z*: [M+H]⁺ calcd for C₁₂H₁₄F₃O₂: 247.0940; Found: 247.0938.

(F) Radical trapping experiments



To a screw-capped vial with stirring bar were added 4CzIPN (3.2 mg, 0.004 mmol, 2 mol%), **1a** (52.8 mg, 0.3 mmol, 1.5 equiv.), Togni II reagent **2c** (63.2 mg, 0.2 mmol, 1.0 equiv.) and TEMPO (125.0 mg, 0.4 mmol, 4 equiv.) in an argon-filled glovebox. After the DMSO (2 mL) was added into the vial *via* a syringe, the vial was sealed with a cap containing a PTFE septum and removed out from the glovebox. The reaction mixture was stirred at room temperature for 48 h

upon irradiation with 24 W blue LED under argon atmosphere. After completion of the reaction, quenched with saturated NaCl (2 mL) and extracted with EtOAc (3 x 5 mL). The combined extracts were dried over MgSO₄, filtered, and evaporated. The analysis of the crude reaction mixture by high resolution mass spectrometry (HRMS) identified compound **7**. **HRMS (ESI)** m/z: [M+H]⁺ calcd for C₁₀H₁₉F₃NO: 226.1413; Found: 226.1421.

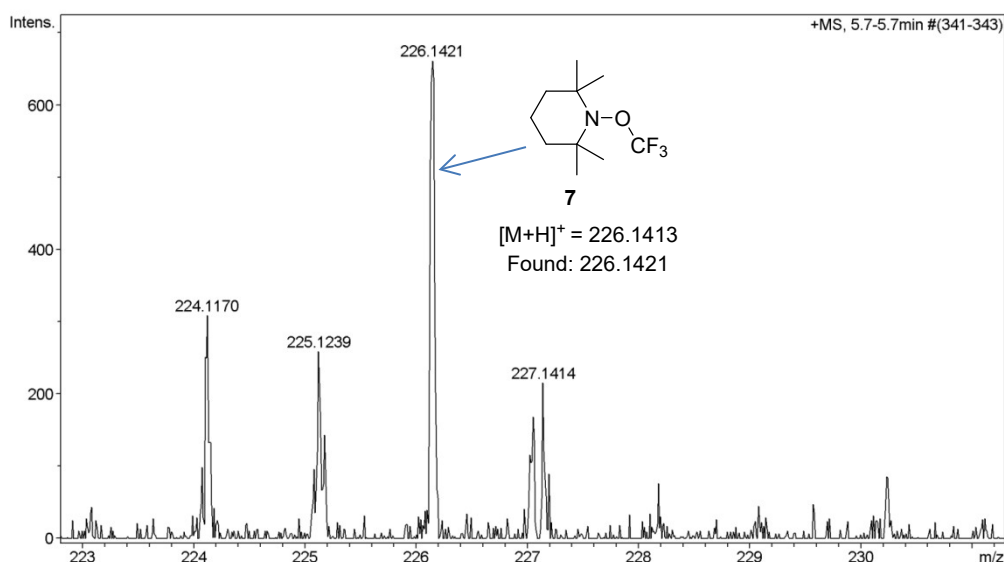


Figure S1. 2-(allyloxy)-3-methylbenzaldehyde **1a** and Togni II reagent **2c** under standard conditions with TEMPO (4.0 equiv.)



To a screw-capped vial with stirring bar were added 4CzIPN (3.2 mg, 0.004 mmol, 2 mol%), **1a** (52.8 mg, 0.3 mmol, 1.5 equiv.), Togni II reagent **2c** (63.2 mg, 0.2 mmol, 1.0 equiv.) and DPE (72.1 mg, 0.4 mmol, 4 equiv.) in an argon-filled glovebox. After the DMSO (2 mL) was added into the vial *via* a syringe, the vial was sealed with a cap containing a PTFE septum and removed out from the glovebox. The reaction mixture was stirred at room temperature for 48 h upon irradiation with 24 W blue LED under argon atmosphere. After completion of the reaction, quenched with saturated NaCl (2 mL) and extracted with EtOAc (3 x 5 mL). The combined extracts were dried over MgSO₄, filtered, and evaporated. The analysis of the crude reaction mixture by high resolution mass spectrometry (HRMS) identified compound **8**. **HRMS (ESI)** m/z: [M+Na]⁺ calcd for C₁₅H₁₁F₃Na: 271.0705; Found: 271.0712.

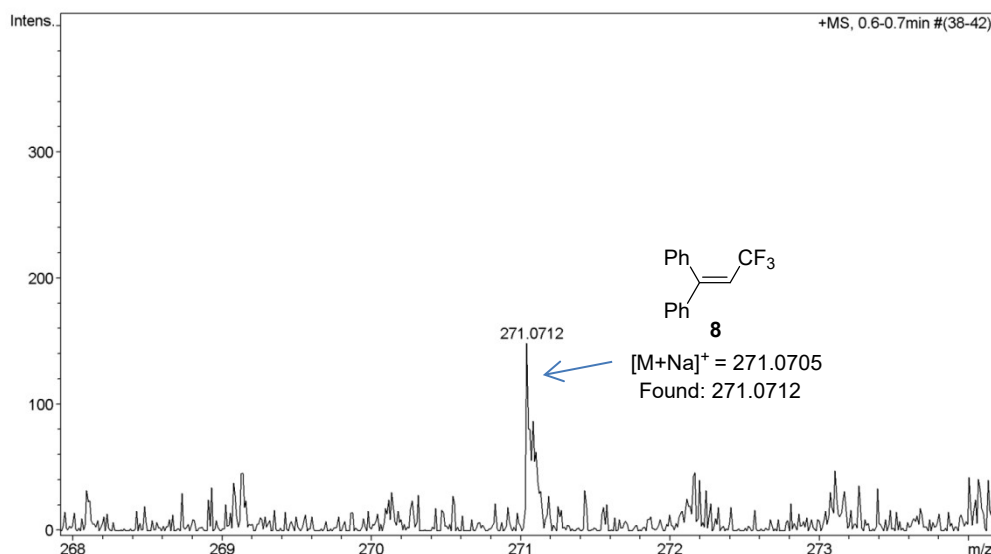


Figure S2. 2-(allyloxy)-3-methylbenzaldehyde **1a** and Togni II reagent **2c** under standard conditions with DPE (4.0 equiv.)

(G) Stern-Volmer fluorescence quenching experiments

Stern-Volmer fluorescence quenching experiments were run with freshly prepared solutions of 1.25 μM 4CzIPN, in anhydrous dry DMSO at room temperature. The solutions were irradiated at 370 nm and fluorescence was measured from 450 nm to 700 nm. Control experiments showed that the excited state 4CzIPN* was mainly quenched by Togni II reagent **2c**.

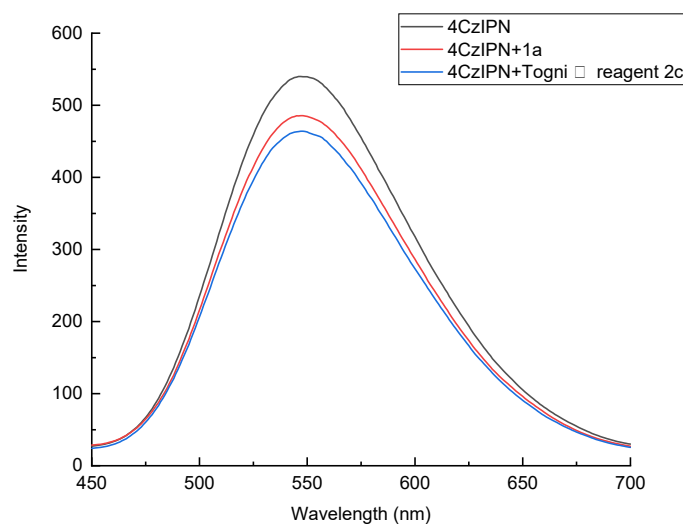


Figure S3. Fluorescence quenching of excited 4CzIPN with **1a**, Togni II reagent **2c** in DMSO (excitation wavelength: 370 nm). 4CzIPN (1.25 μM) in DMSO (black line), 4CzIPN (1.25 μM) with **1a** (5.0 mM) in DMSO (red line), 4CzIPN (1.25 μM) with Togni II reagent **2c** (5.0 mM) in DMSO (blue line).

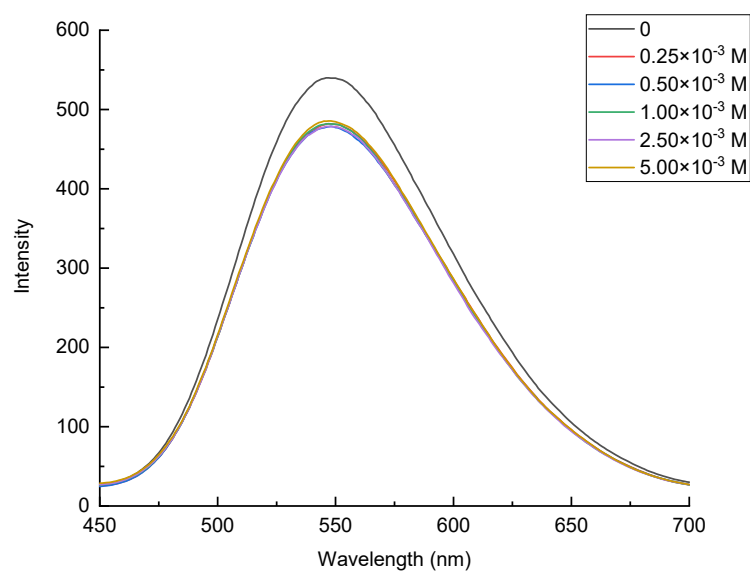


Figure S4. Luminescence quenching of 4CzIPN by **1a**

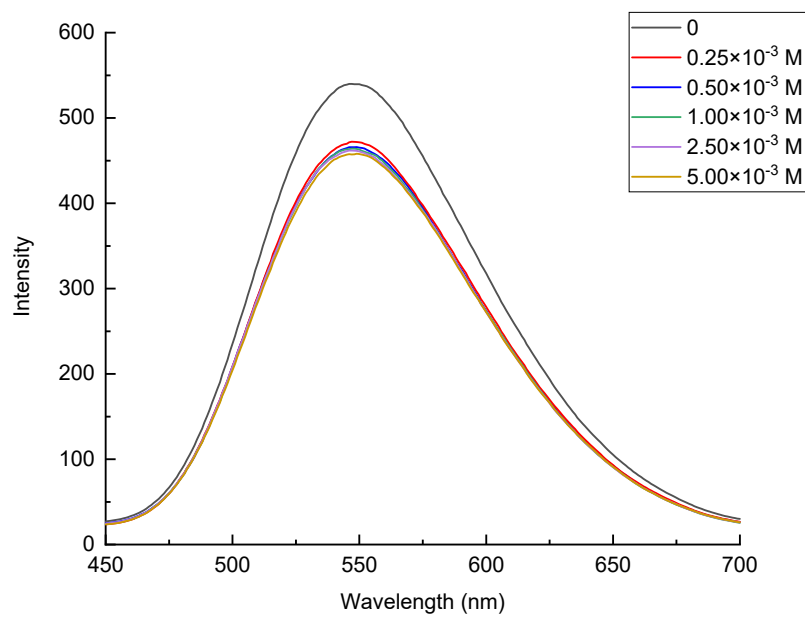


Figure S5. Luminescence quenching of 4CzIPN by Togni II reagent **2c**

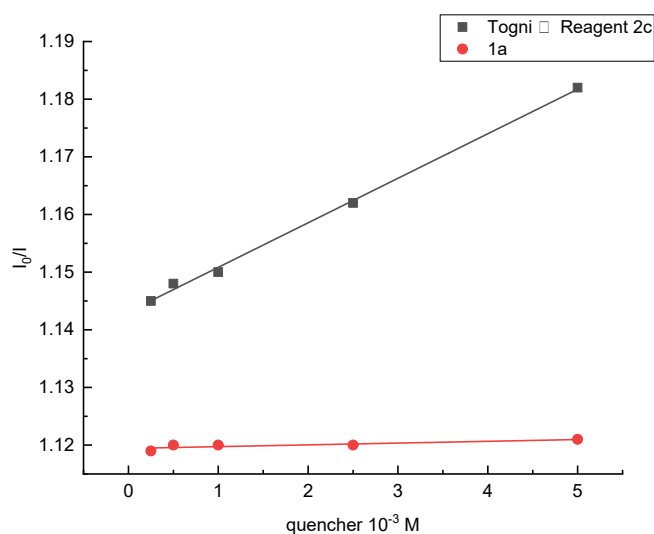


Figure S6 Stern-Volmer plots of 4CzIPN and two quenchers. I_0 and I are luminescence intensities in the absence and presence of the indicated concentrations (10^{-3} M) of the corresponding quencher.

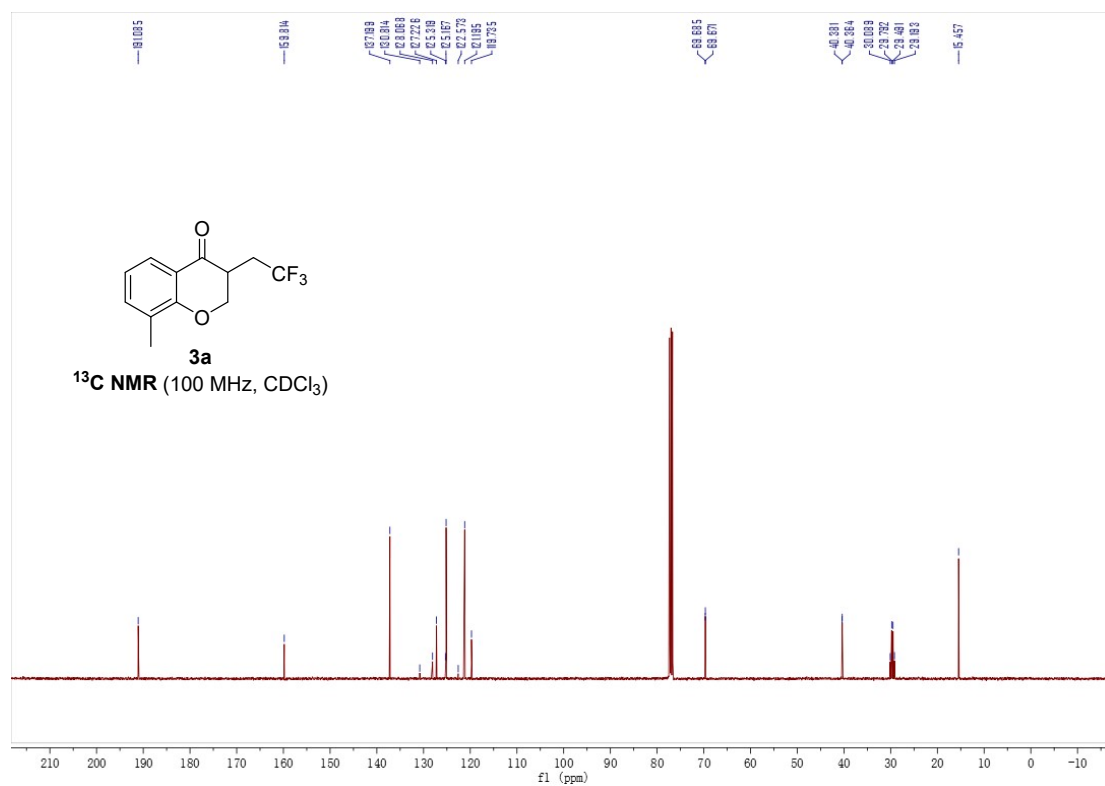
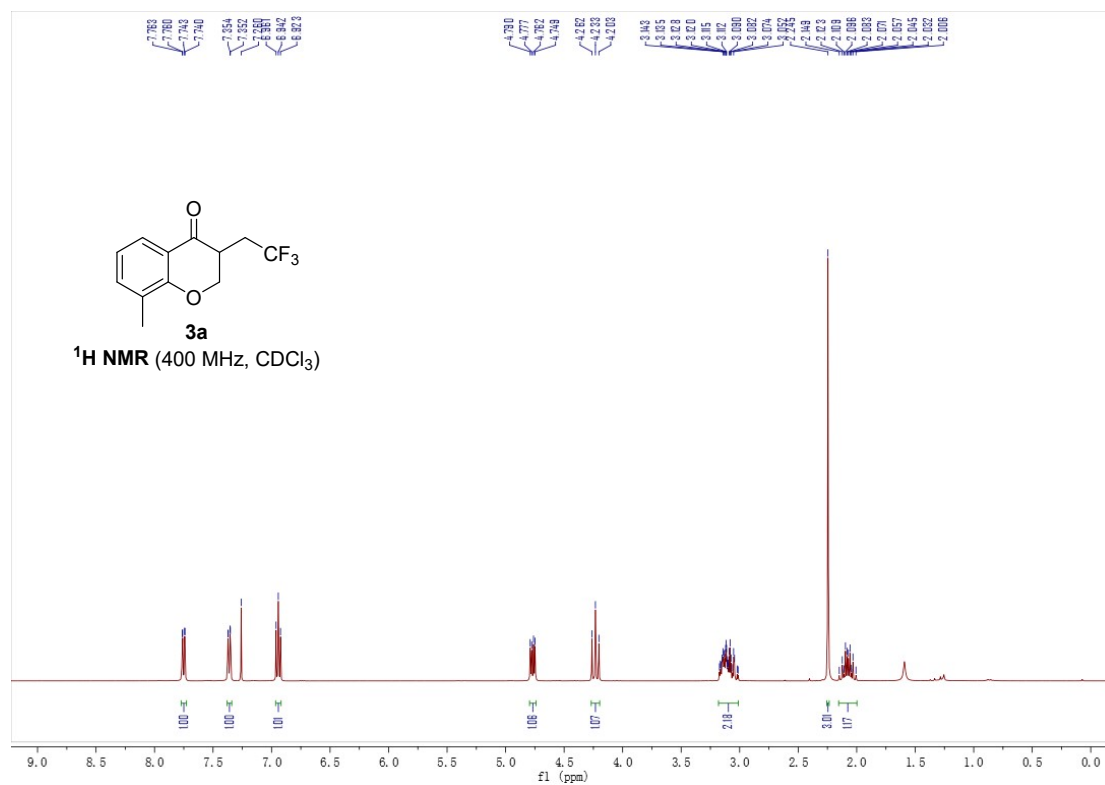
These results suggested that the excited photocatalyst 4CzIPN was primarily quenched by Togni II reagent **2c**, while **1a** showed much less effect.

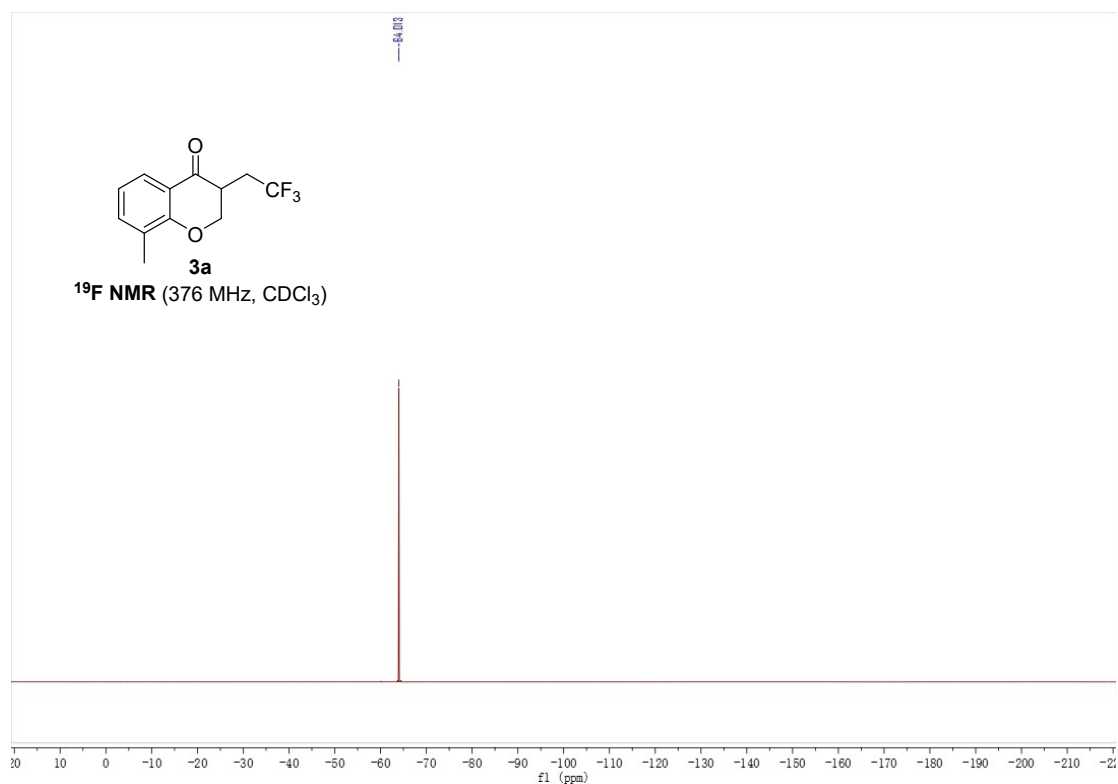
References:

1. Zhao, J.; Li, P.; Li, X.; Xia, C.; Li, F. Straightforward Synthesis of Functionalized Chroman-4-ones through Cascade Radical Cyclization-Coupling of 2-(allyloxy)arylaldehydes. *Chem. Commu.* **2016**, 52, 3661-3664.
2. Li, D.-Y.; Wei, Y.; Marek, I.; Tang, X.-Y.; Shi, M. Gold(I)-catalyzed Cycloisomerization of Vinylidenecyclopropane-enes via Carbene or Non-carbene Processes. *Chem. Sci.* **2015**, 6, 5519-5525.
3. Che, J.; Ma, C.; Lu, J.; Chen, B.; Shi, Q.; Jin, X.; Song, R.; Xu, F.; Gan, L.; Li, J.; Hu, Y.; Dong, X. Discovery of Seneciobipyrrolidine Derivatives for the Amelioration of Glucose Homeostasis Disorders through 4E-BP1/Akt/AMPK Signaling Activation. *Eur. J. Med. Chem.* **2022**, 228.
4. Hu, H.; Chen, X.; Sun, K.; Wang, J.; Liu, Y.; Liu, H.; Fan, L.; Yu, B.; Sun, Y.; Qu, L.; Zhao, Y. Silver-Catalyzed Radical Cascade Cyclization toward 1,5-/1,3-Dicarbonyl Heterocycles: An Atom-/Step-Economical Strategy Leading to Chromenopyridines and Isoxazole-/Pyrazole-Containing Chroman-4-Ones. *Org. Lett.* **2018**, 20, 6157-6160.
5. Gao, F.; Meng, F.-X.; Du, J.-Y.; Zhang, S.; Huang, H.-L. One-Step Synthesis of Trifluoroethylated Chromones via Radical Cascade Cyclization-Coupling of 2-(Allyloxy)arylaldehydes. *Eur. J. Org. Chem.* **2020**, 2020, 209-212.

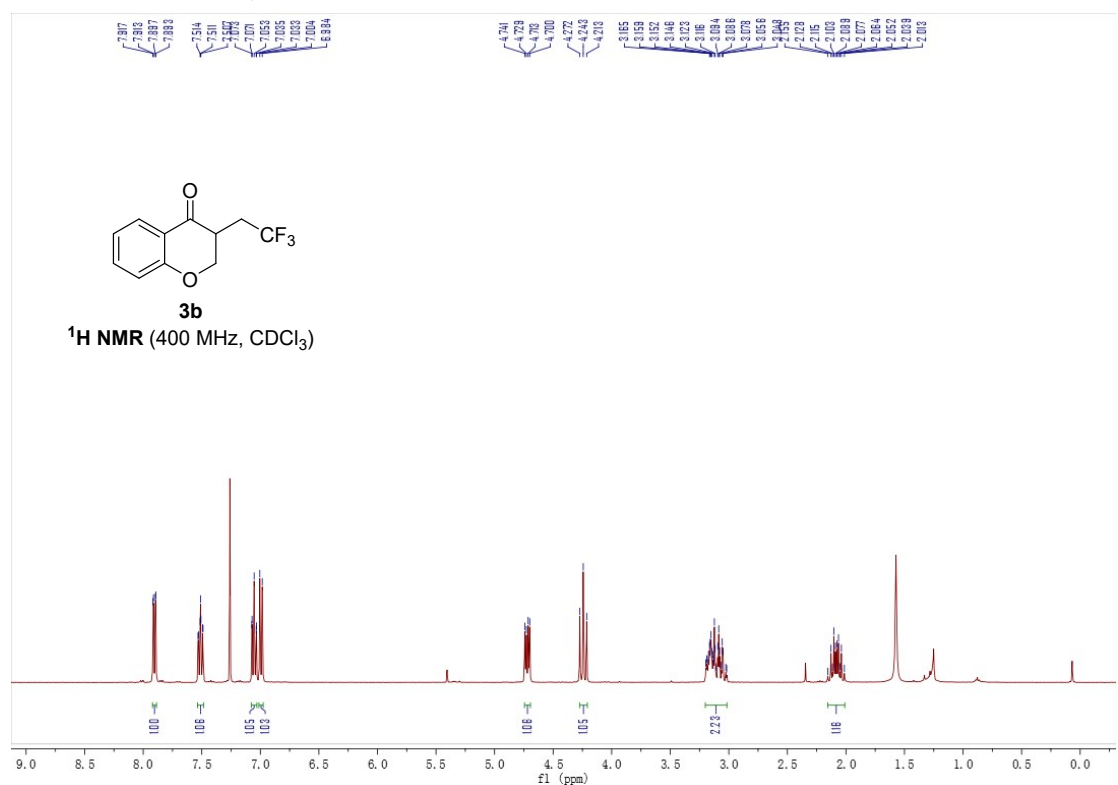
(H) NMR spectra of new compounds

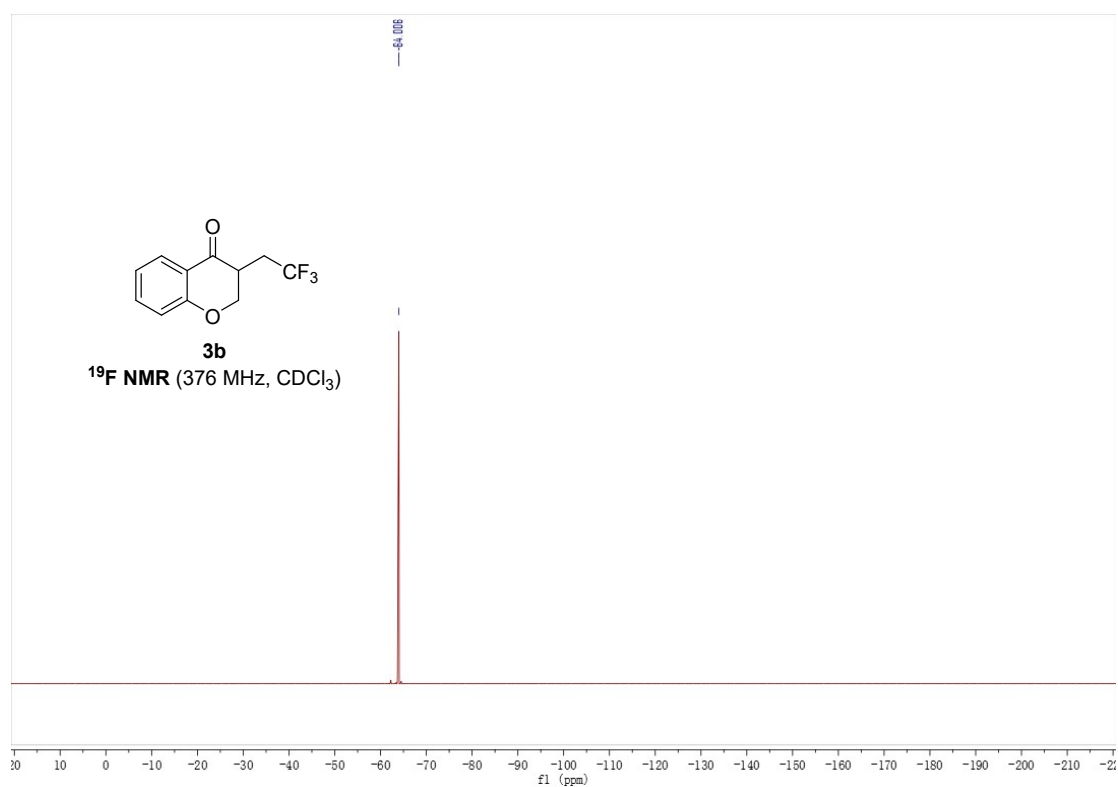
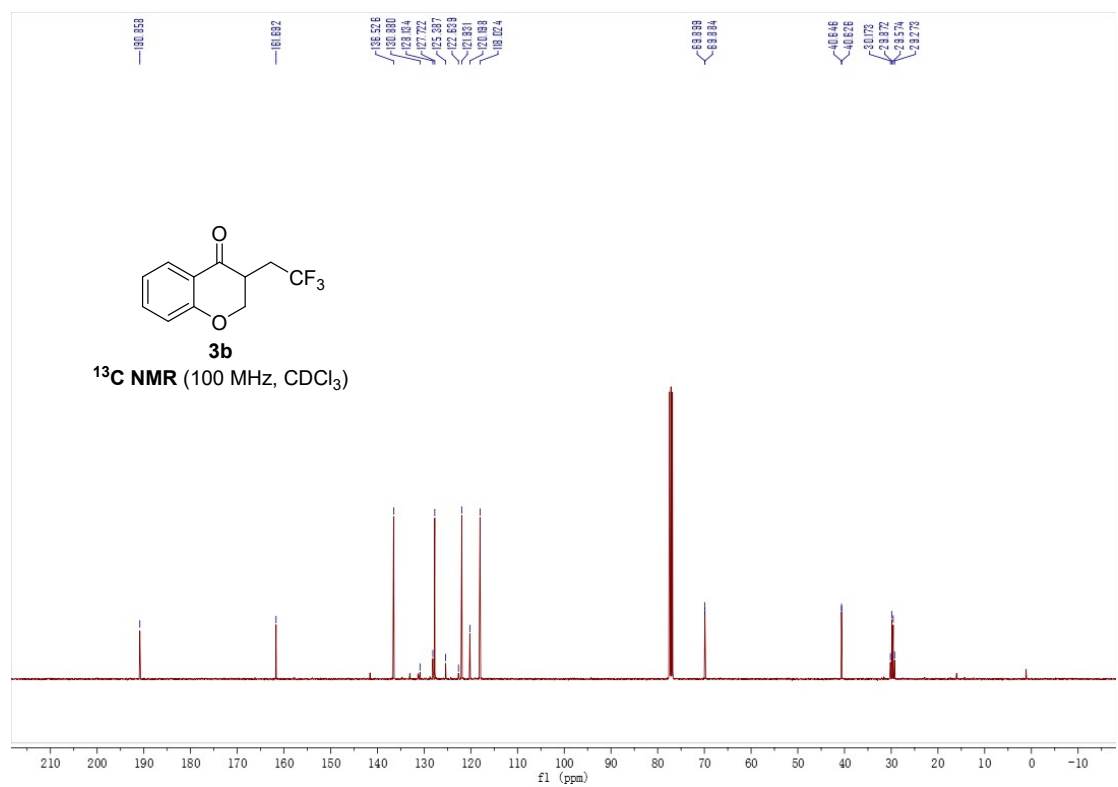
(S)-8-Methyl-3-(2,2,2-trifluoroethyl)chroman-4-one (3a)



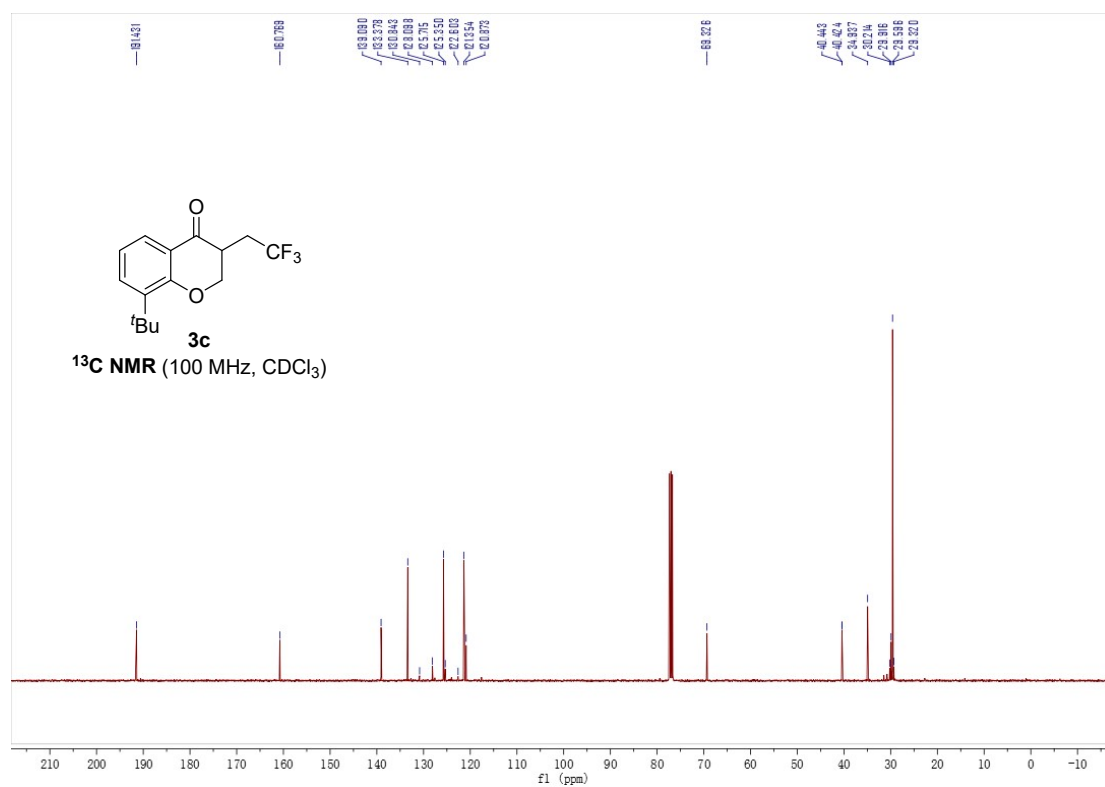
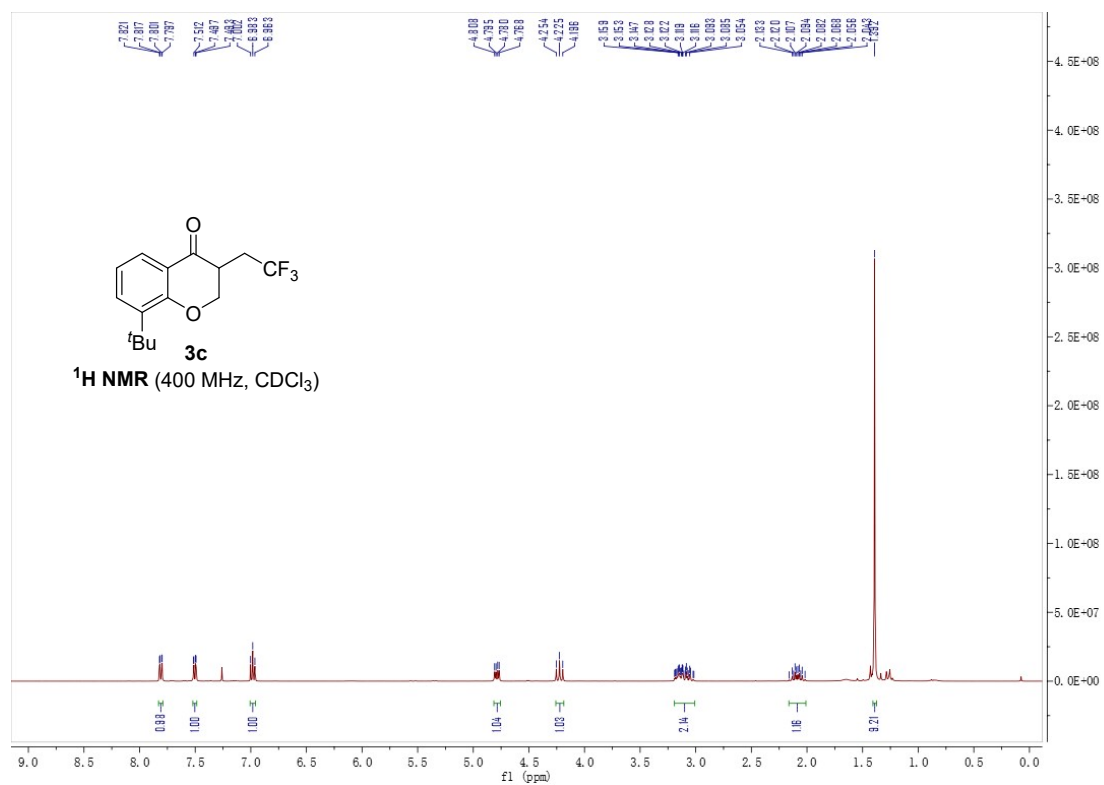


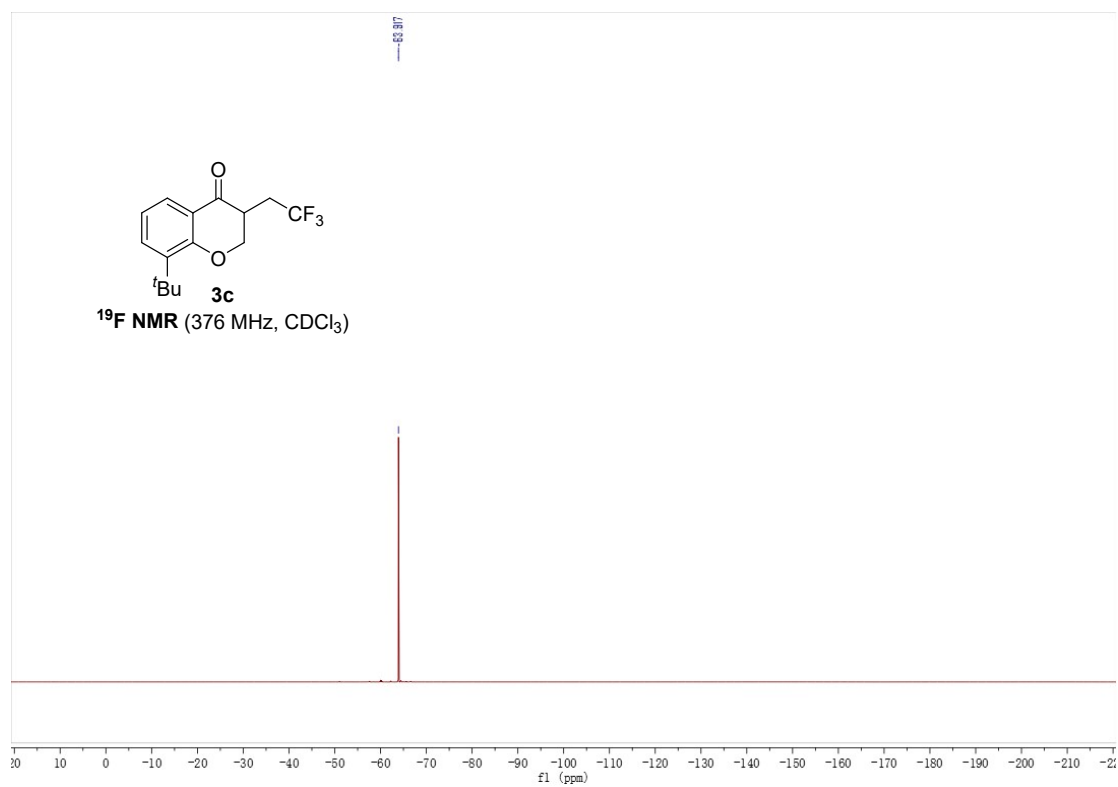
3-(2,2,2-trifluoroethyl)chroman-4-one (3b)



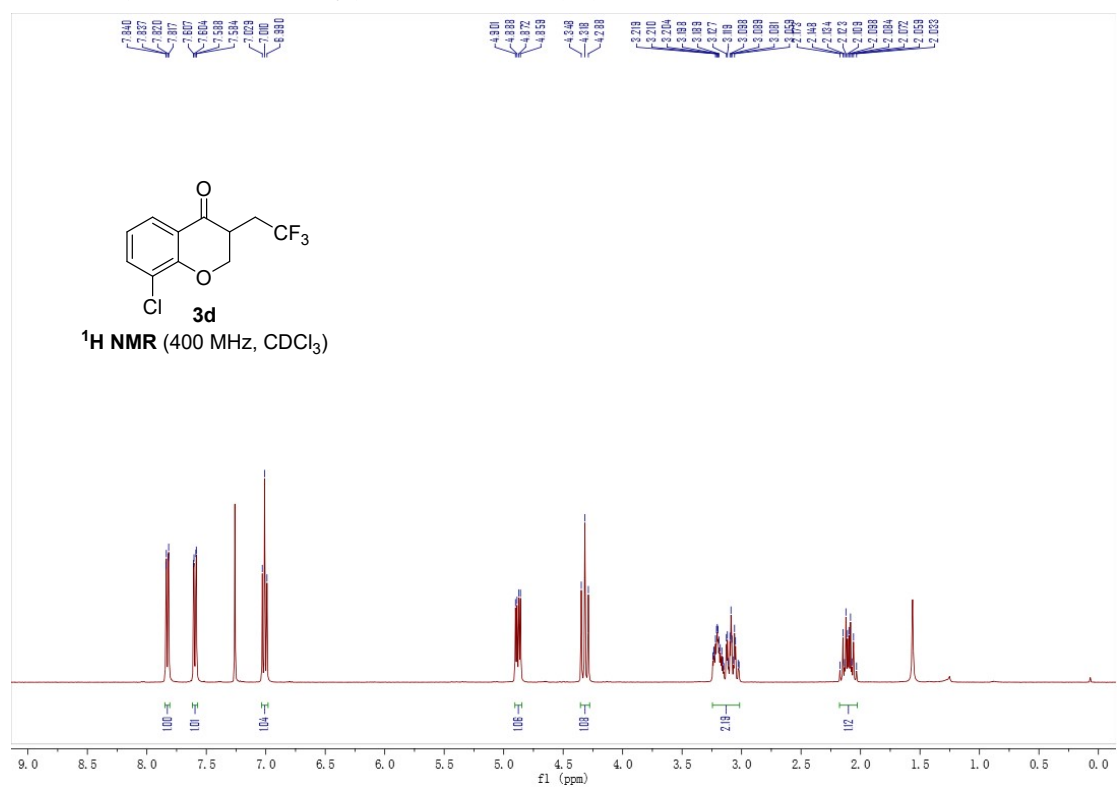


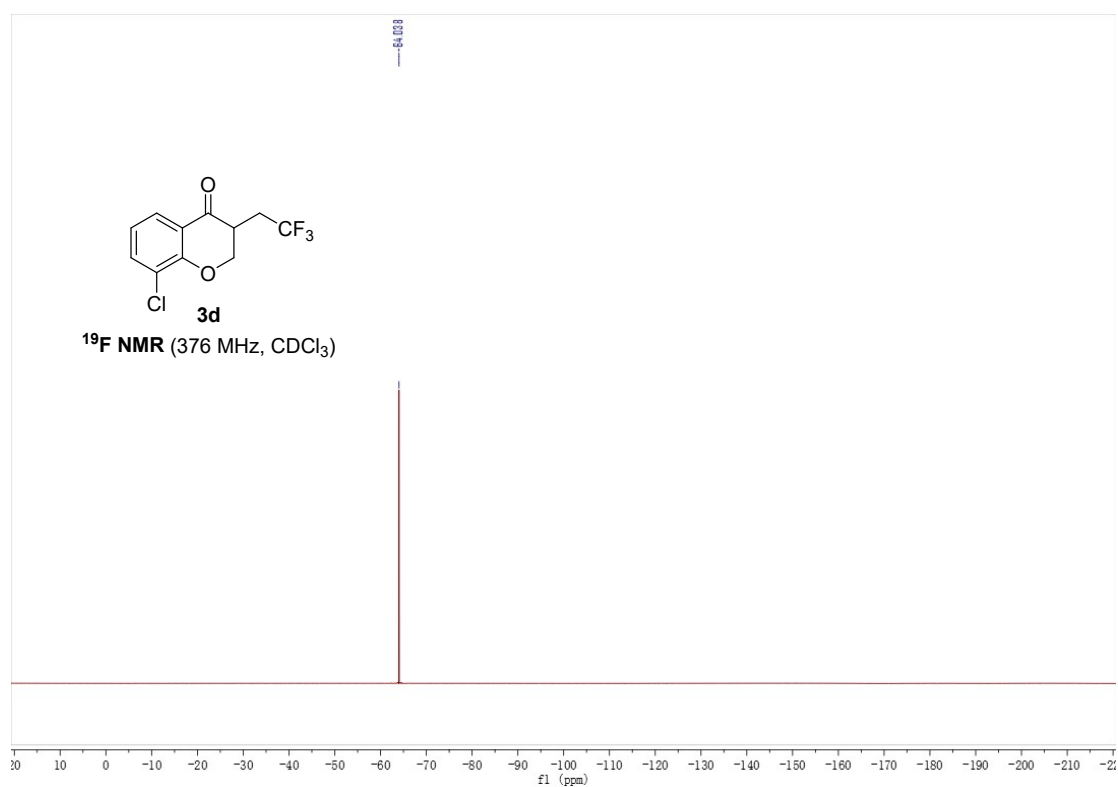
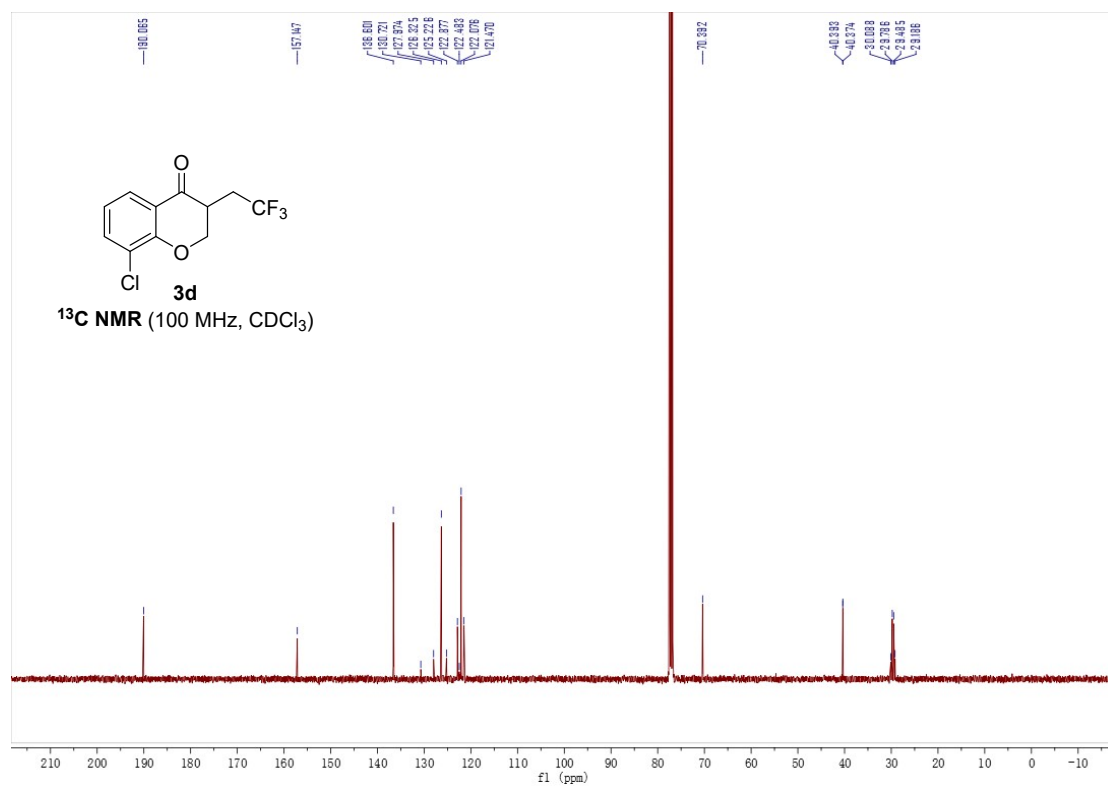
8-(tert-butyl)-3-(2,2,2-trifluoroethyl)chroman-4-one (3c)



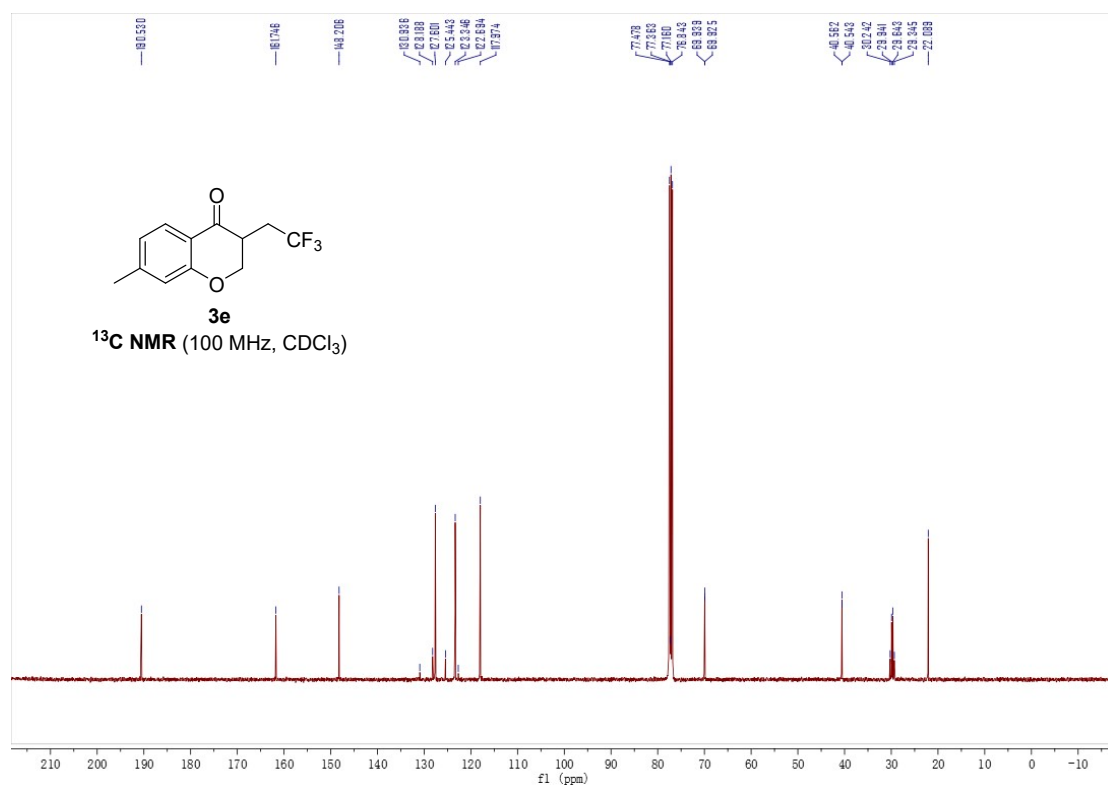
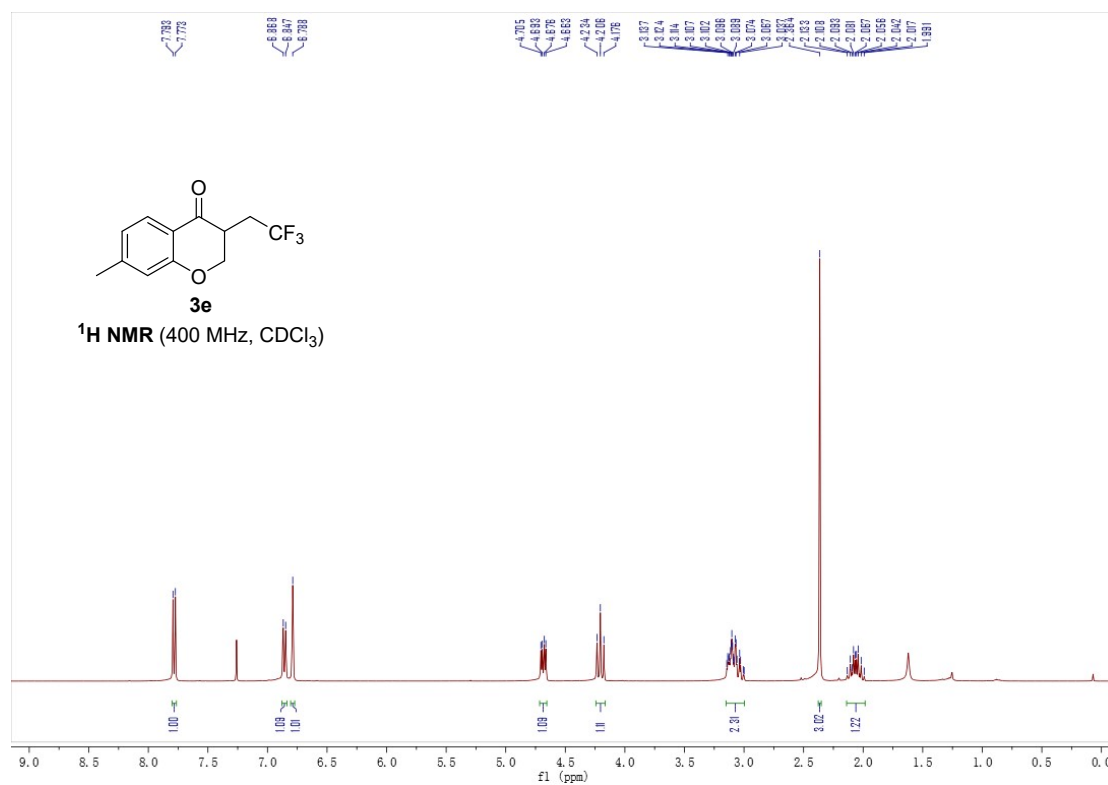


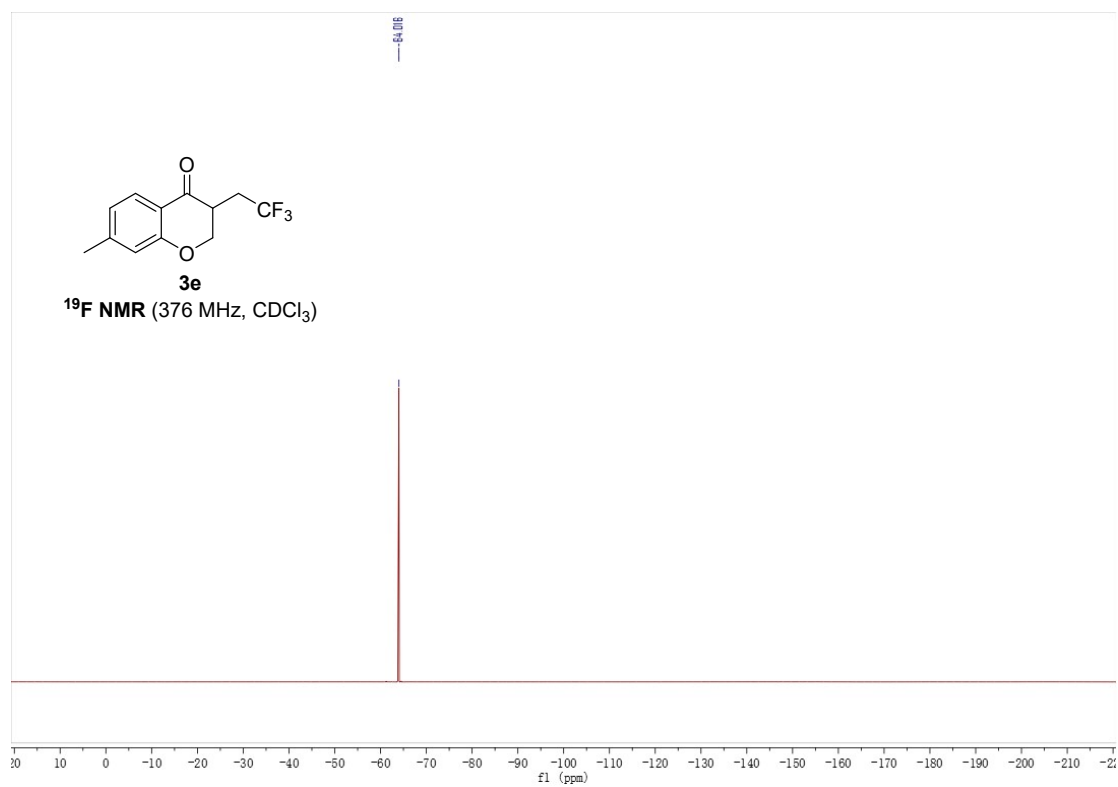
chloro-3-(2,2,2-trifluoroethyl)chroman-4-one (3d)



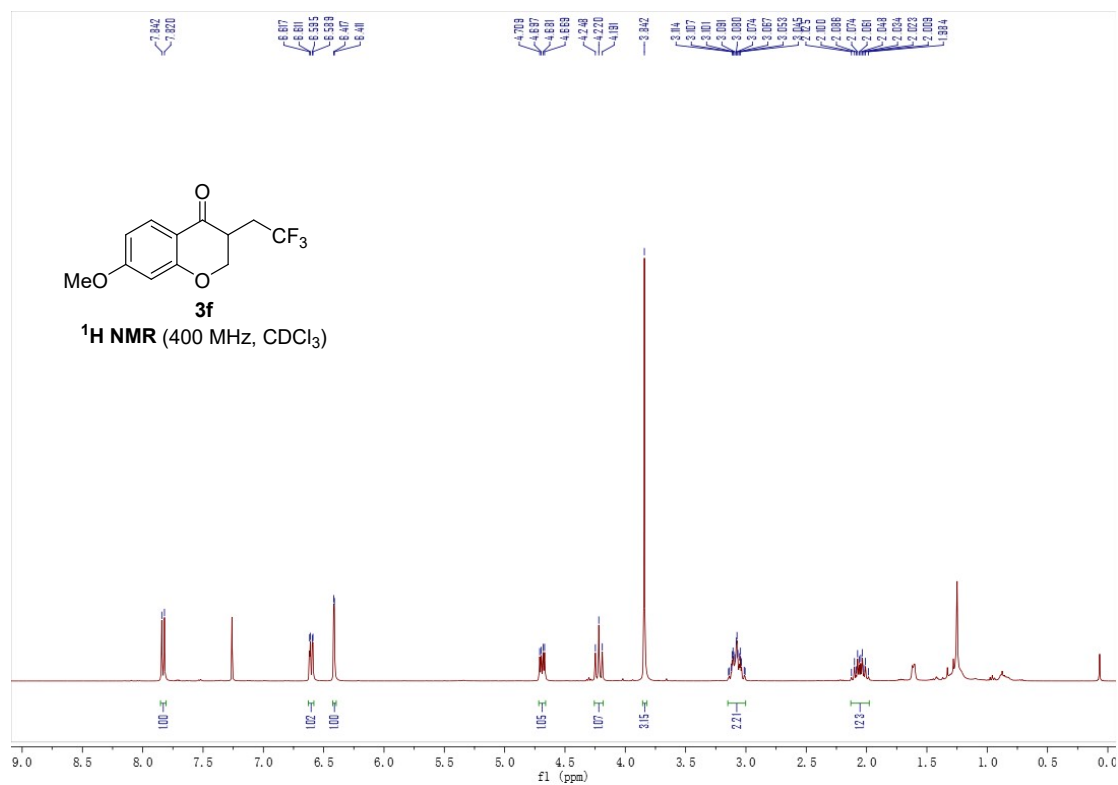


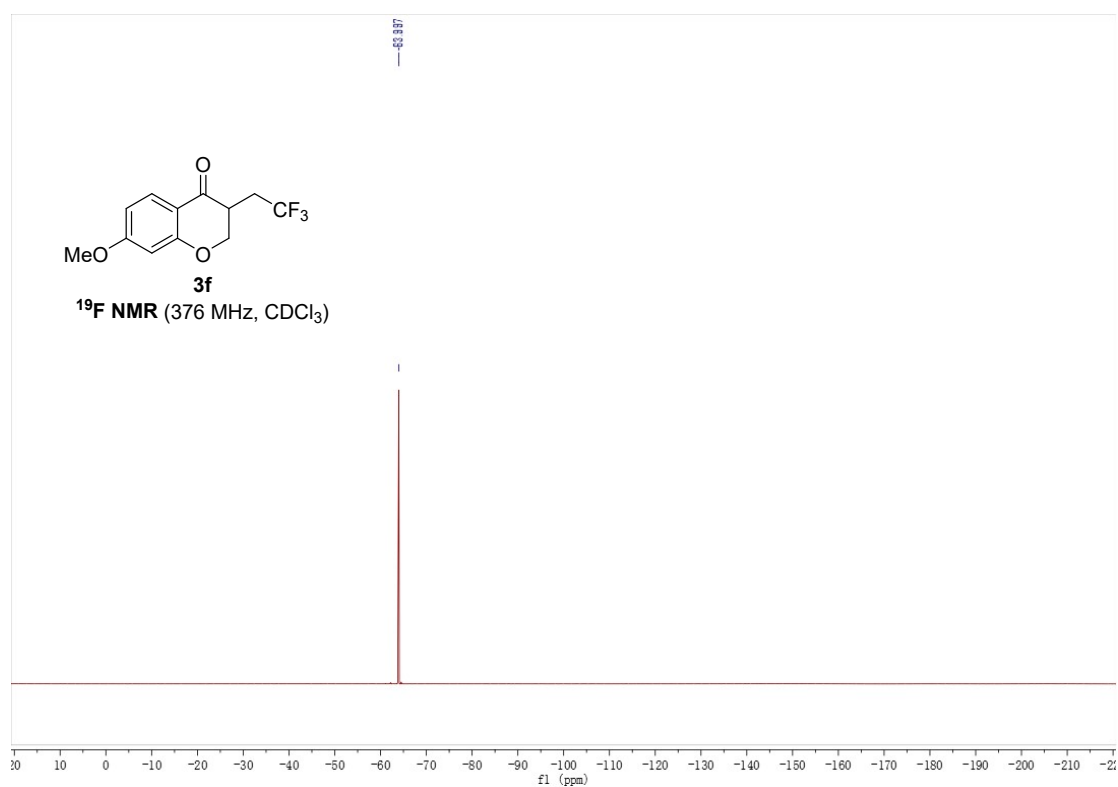
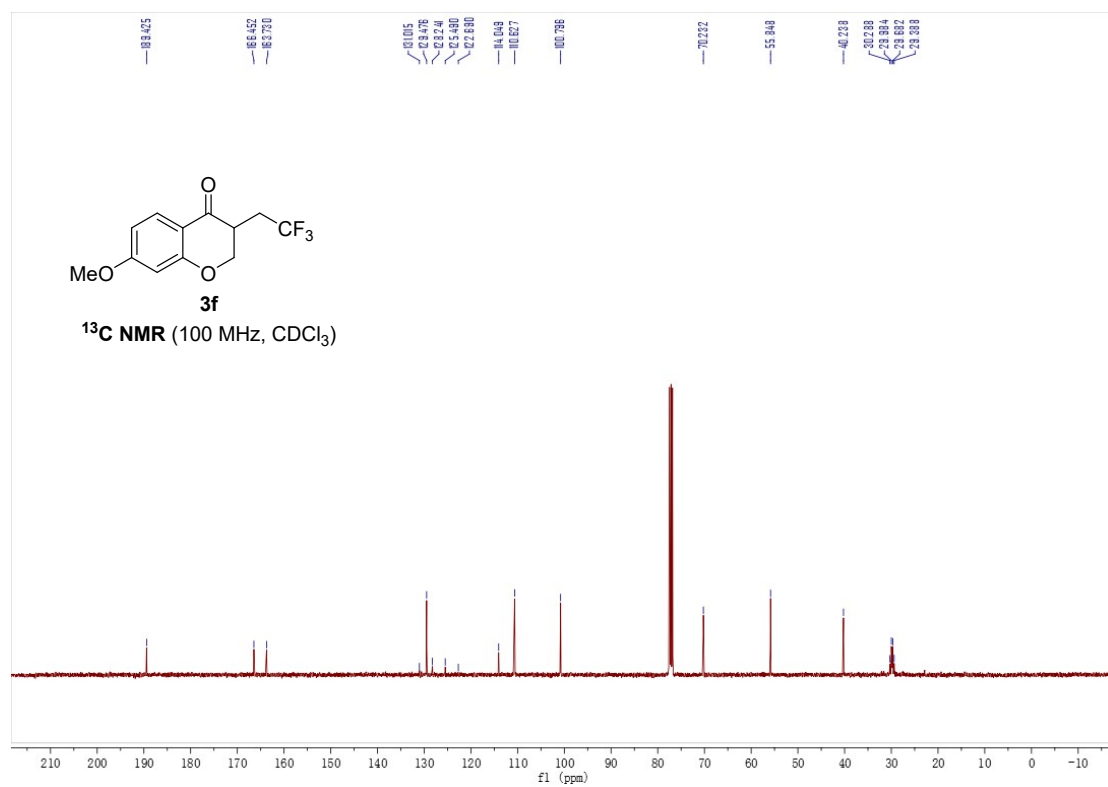
7-methyl-3-(2,2,2-trifluoroethyl)chroman-4-one (3e)



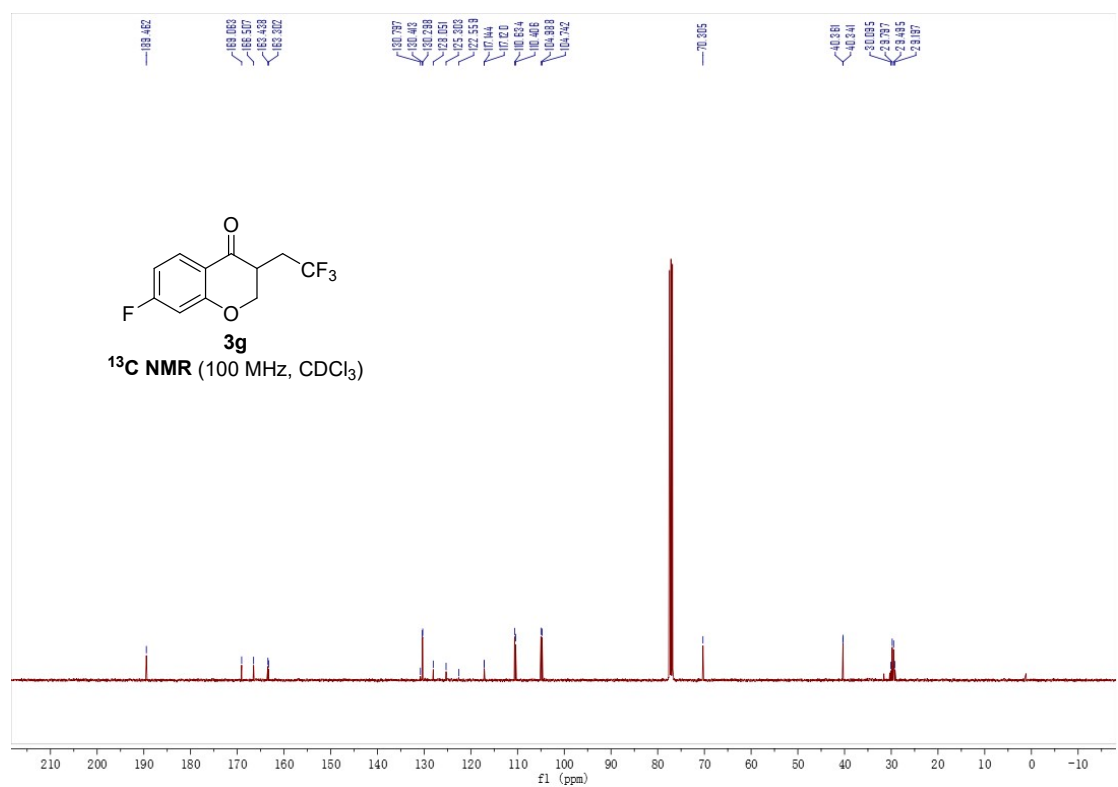
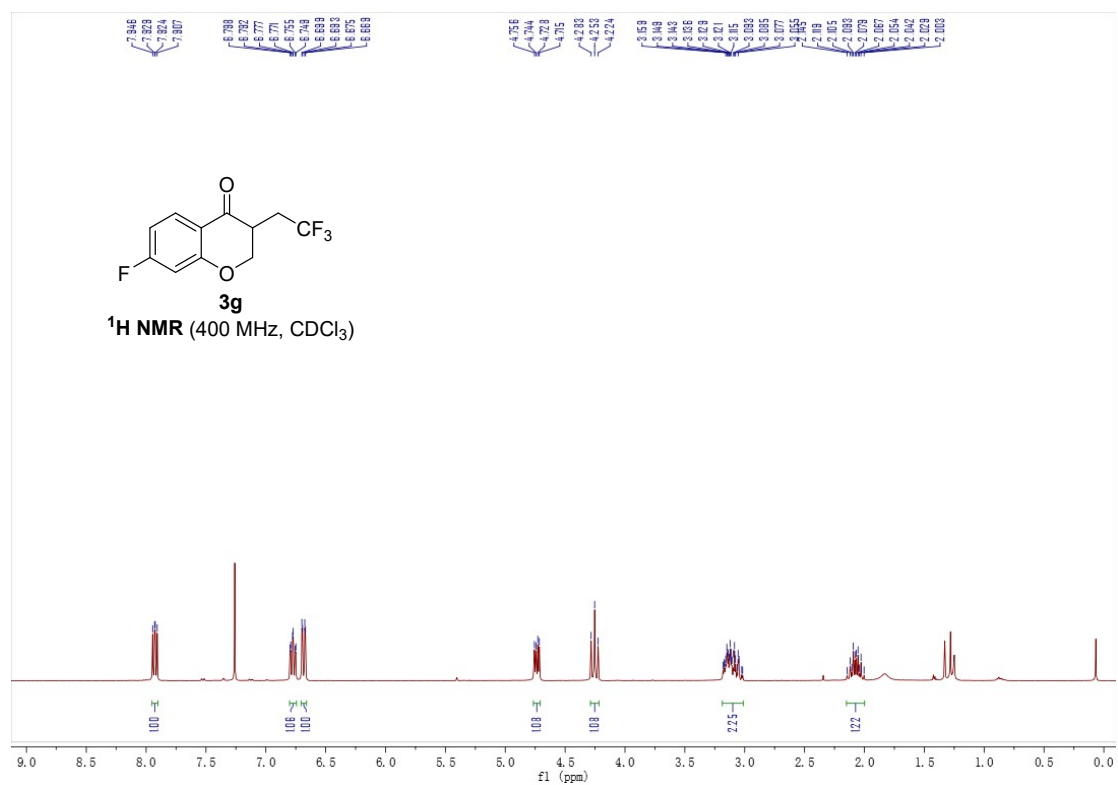


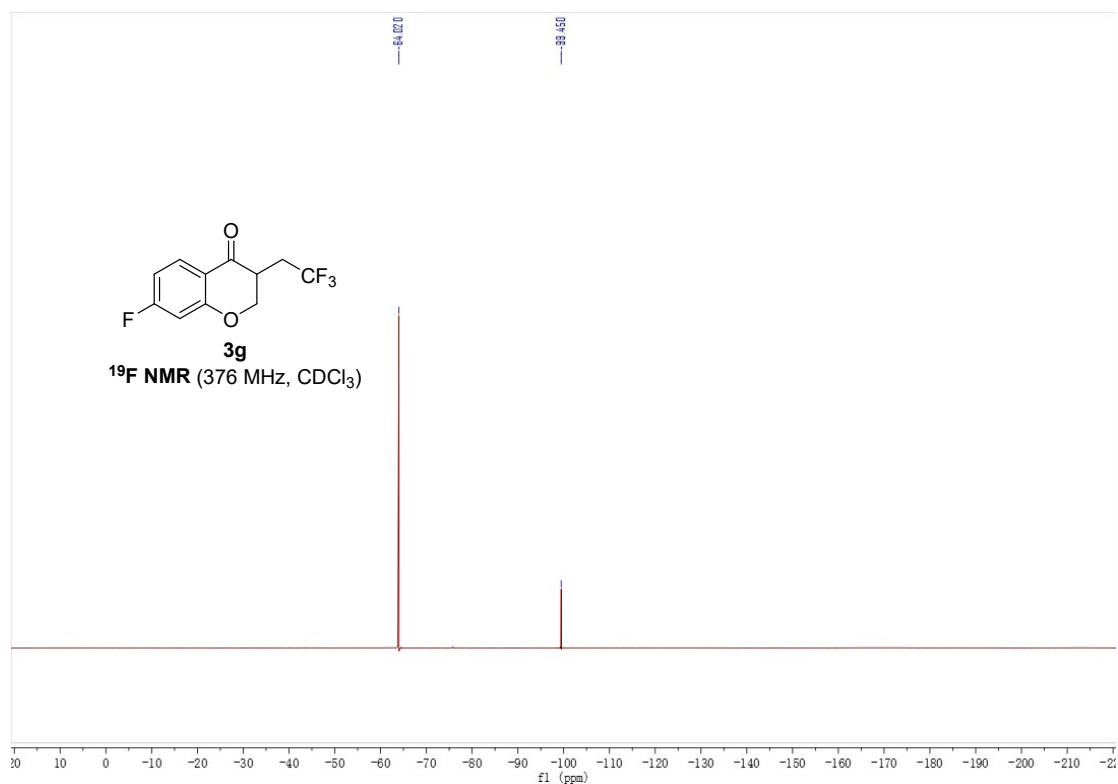
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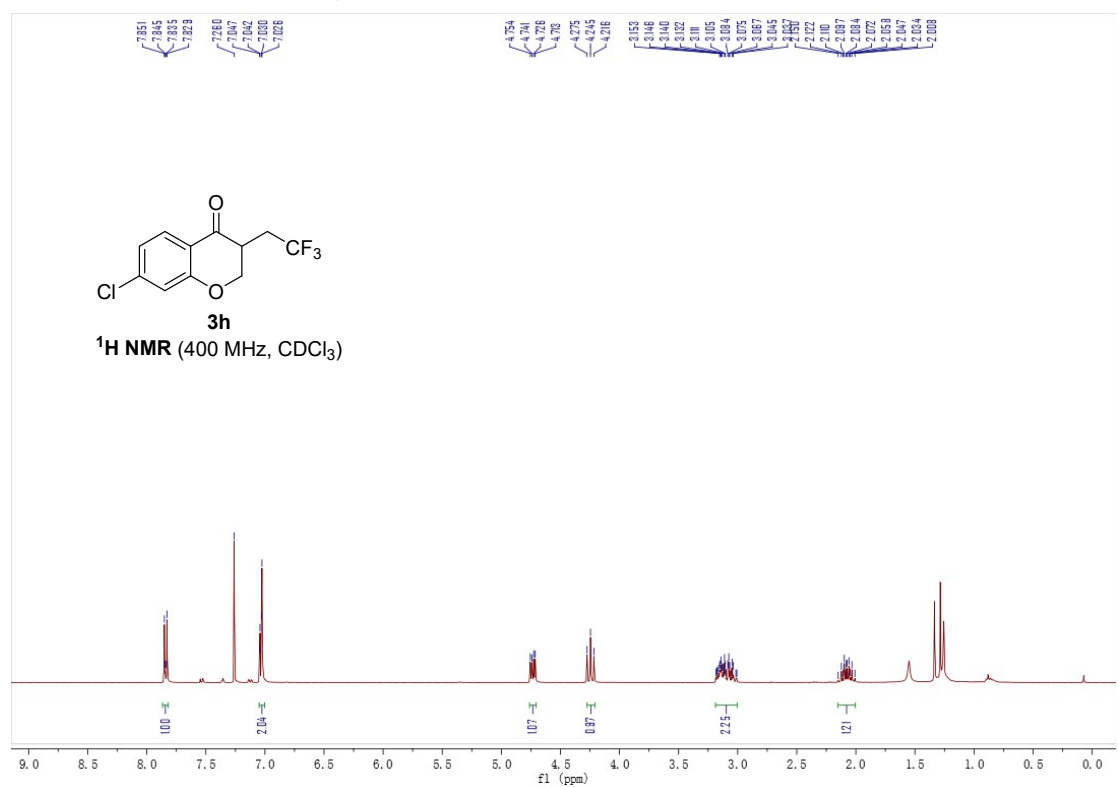


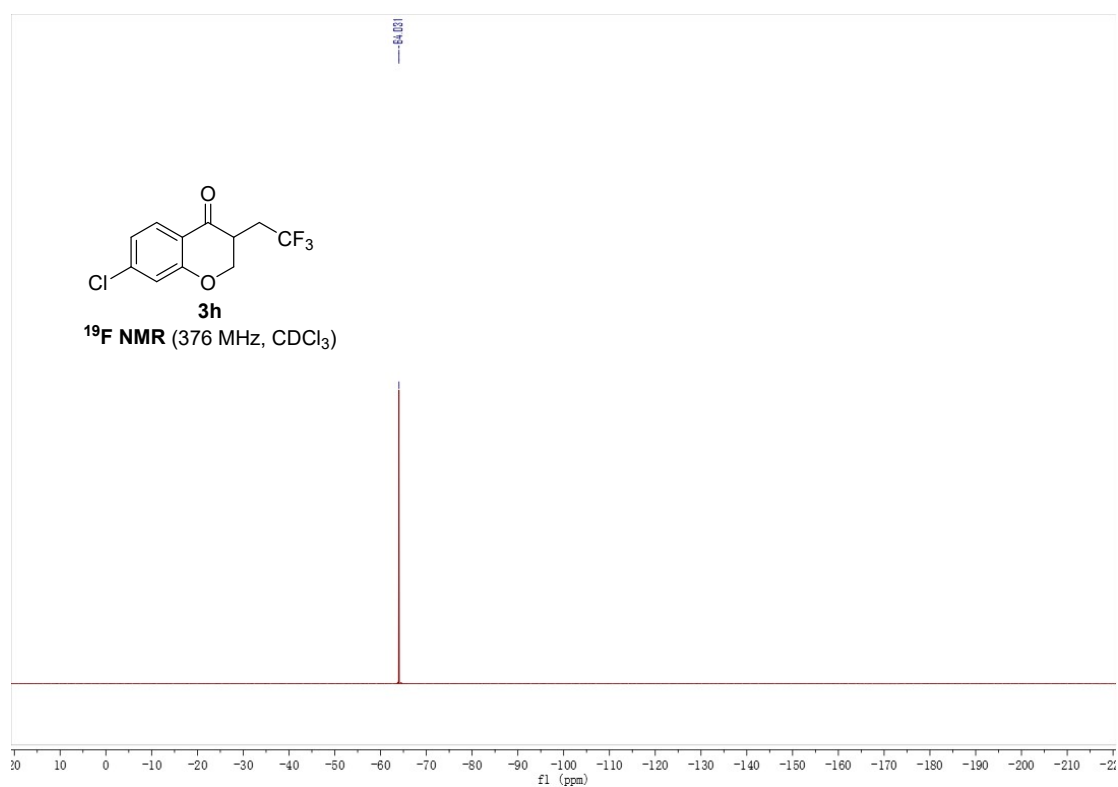
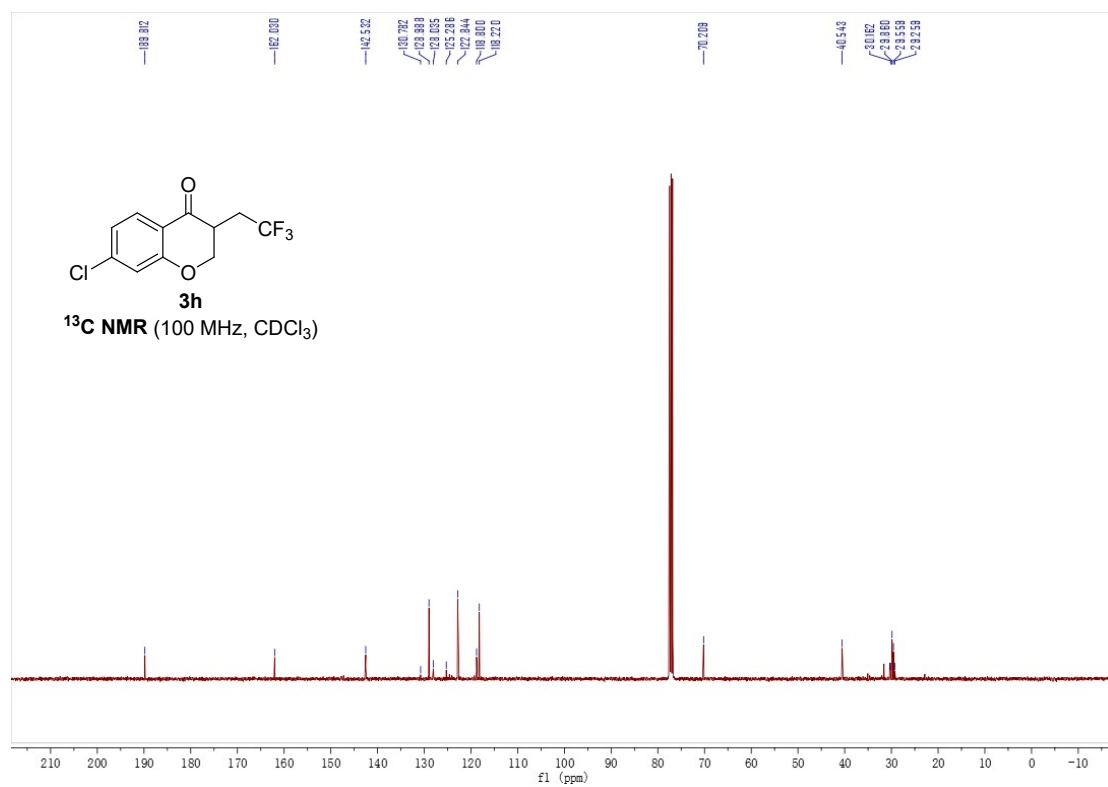
7-fluoro-3-(2,2,2-trifluoroethyl)chroman-4-one (3g)



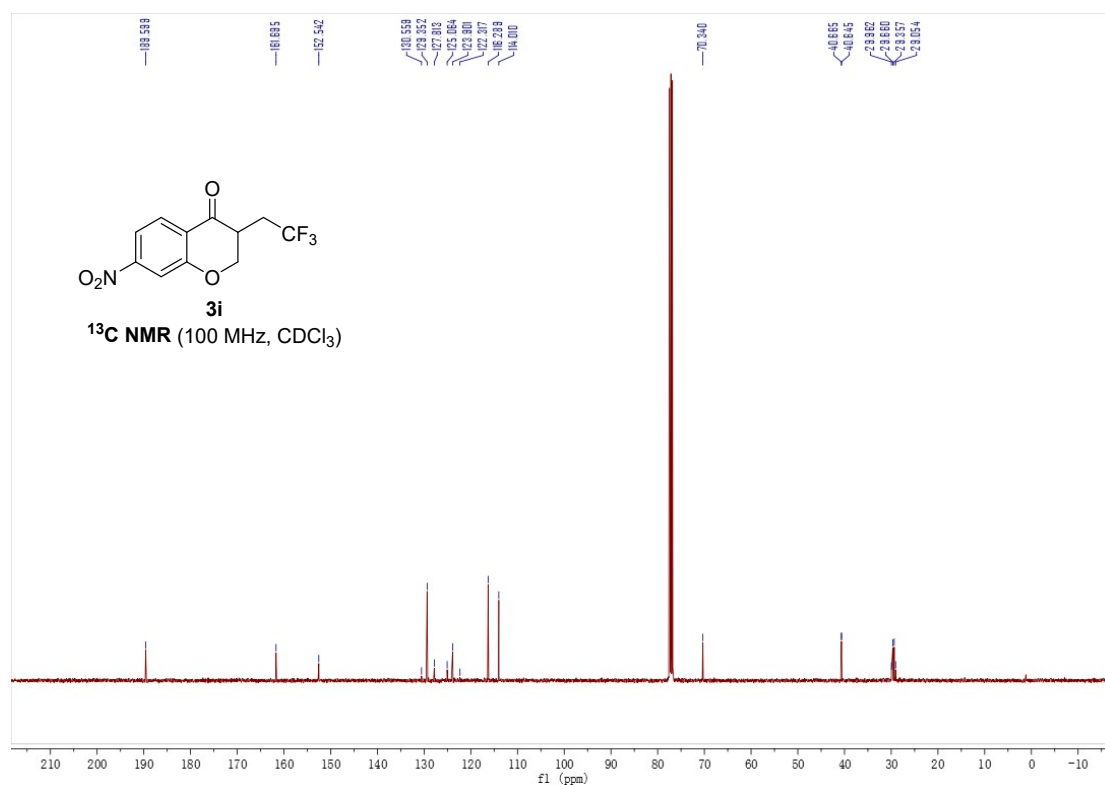
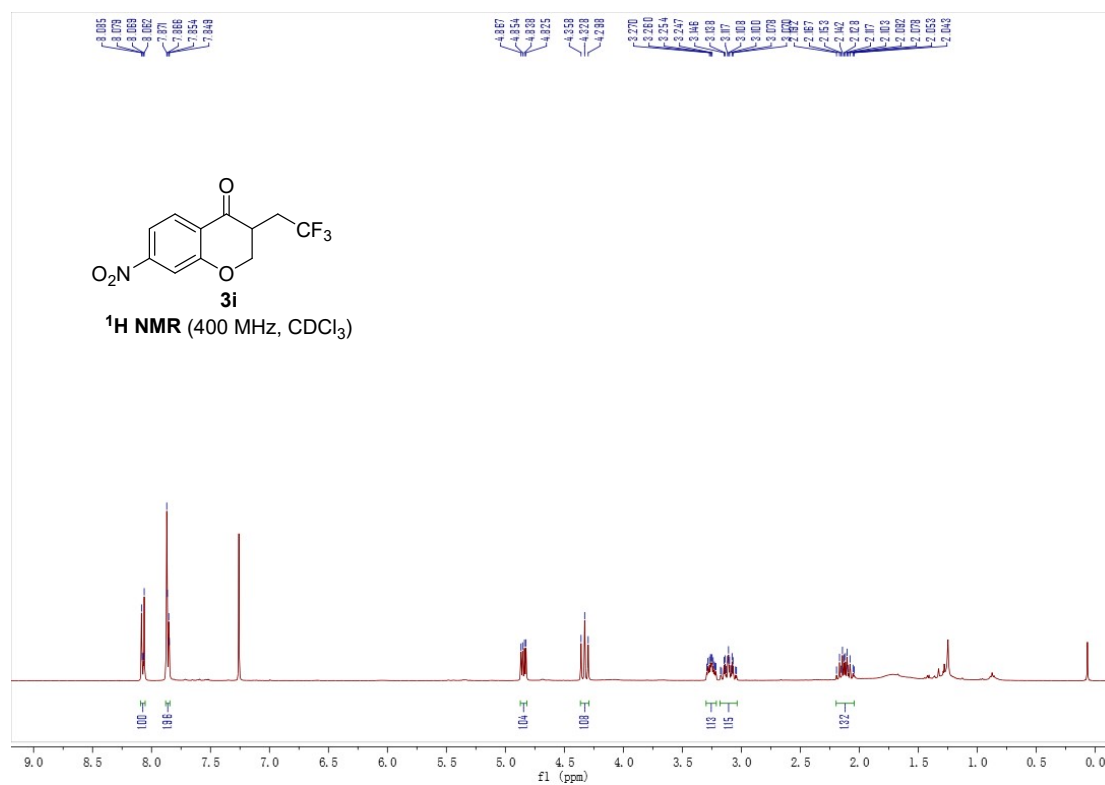


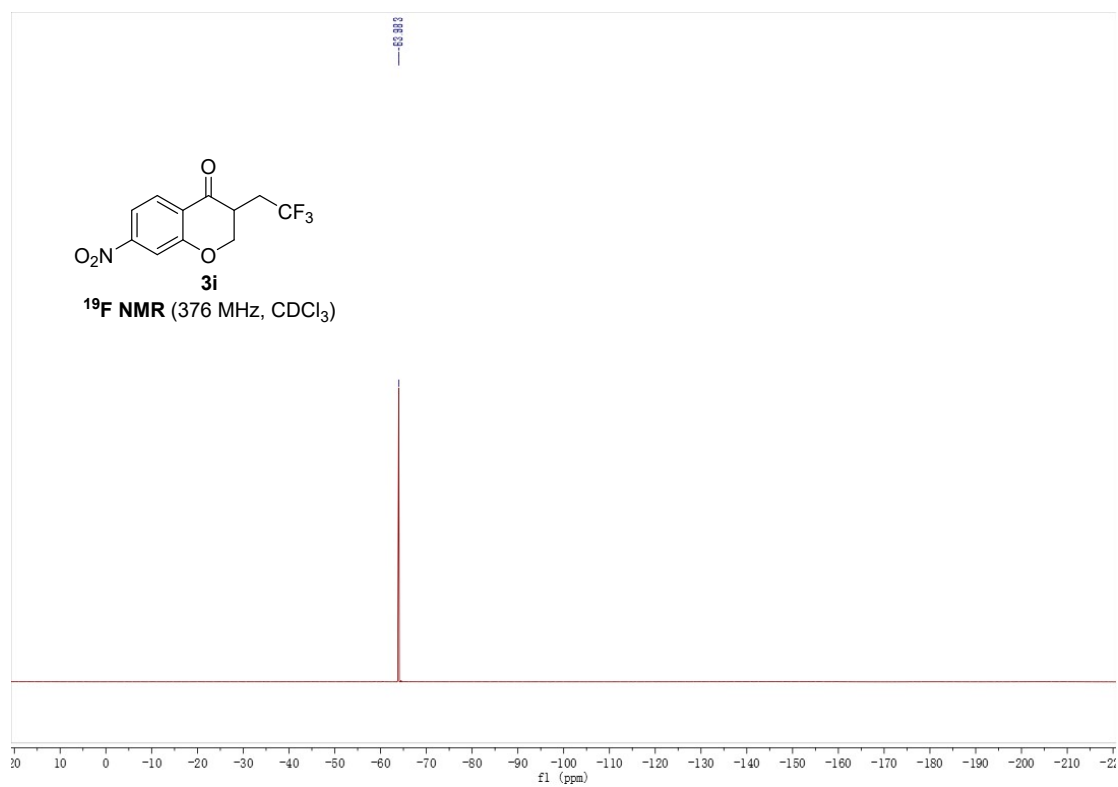
chloro-3-(2,2,2-trifluoroethyl)chroman-4-one (3h)



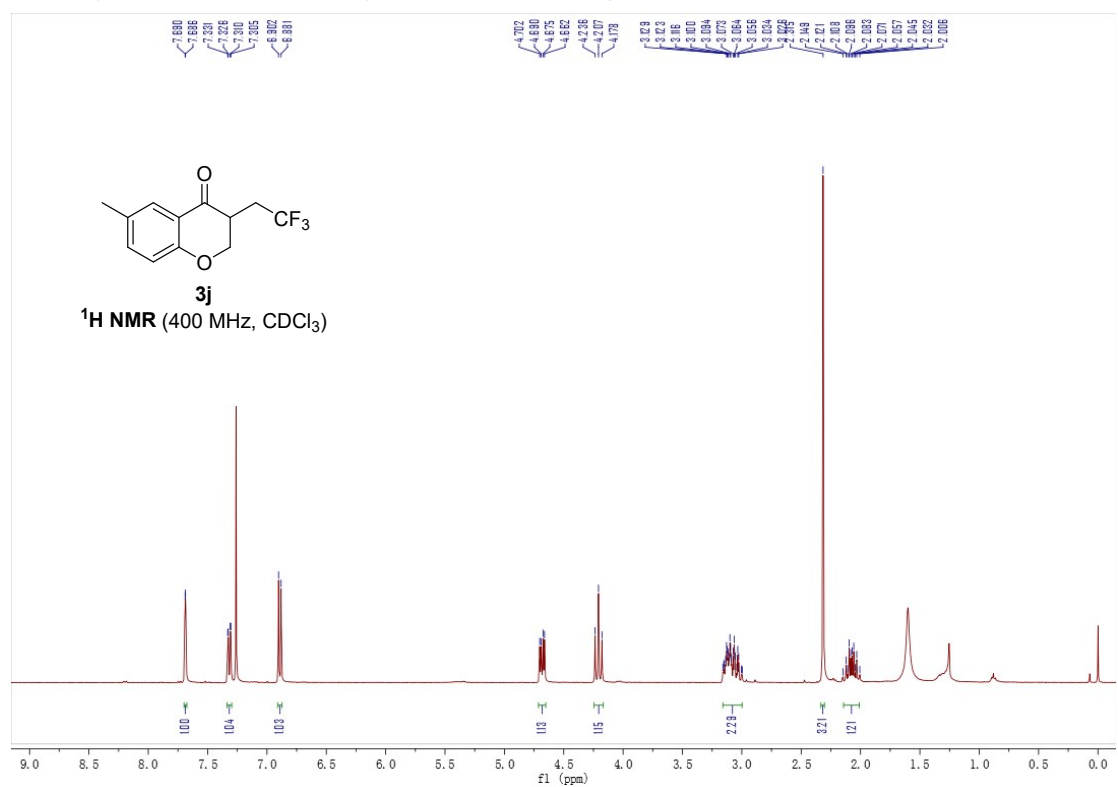


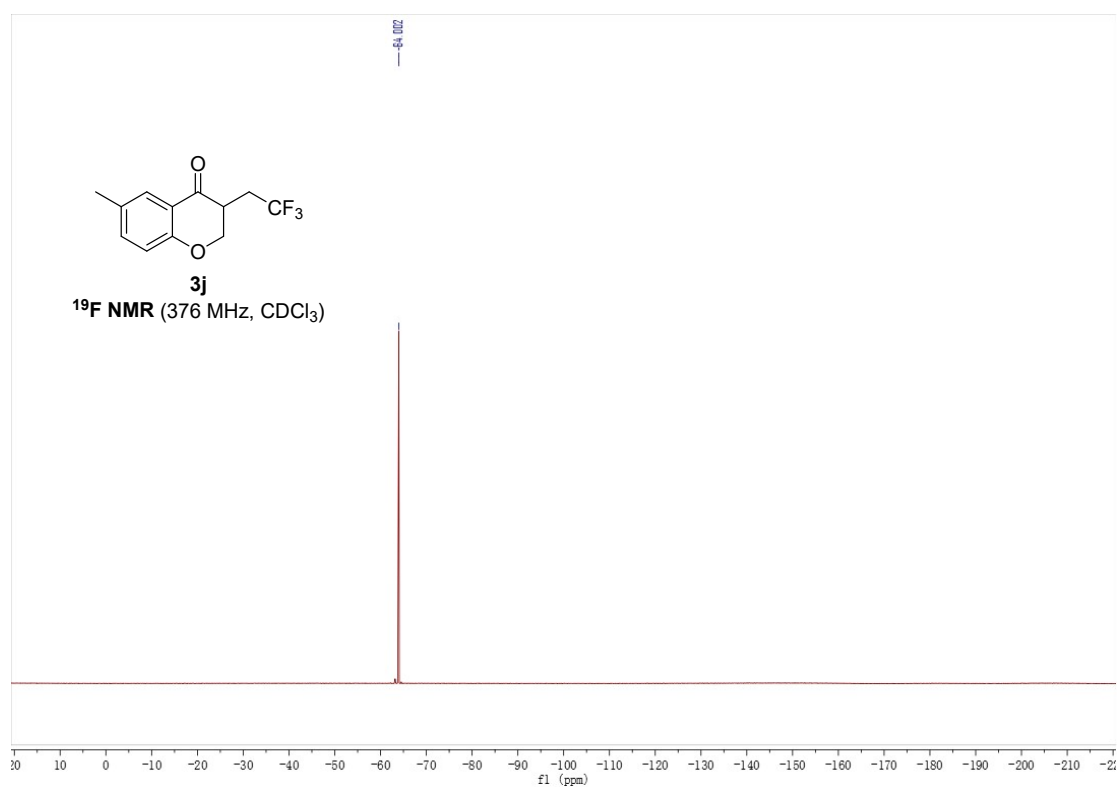
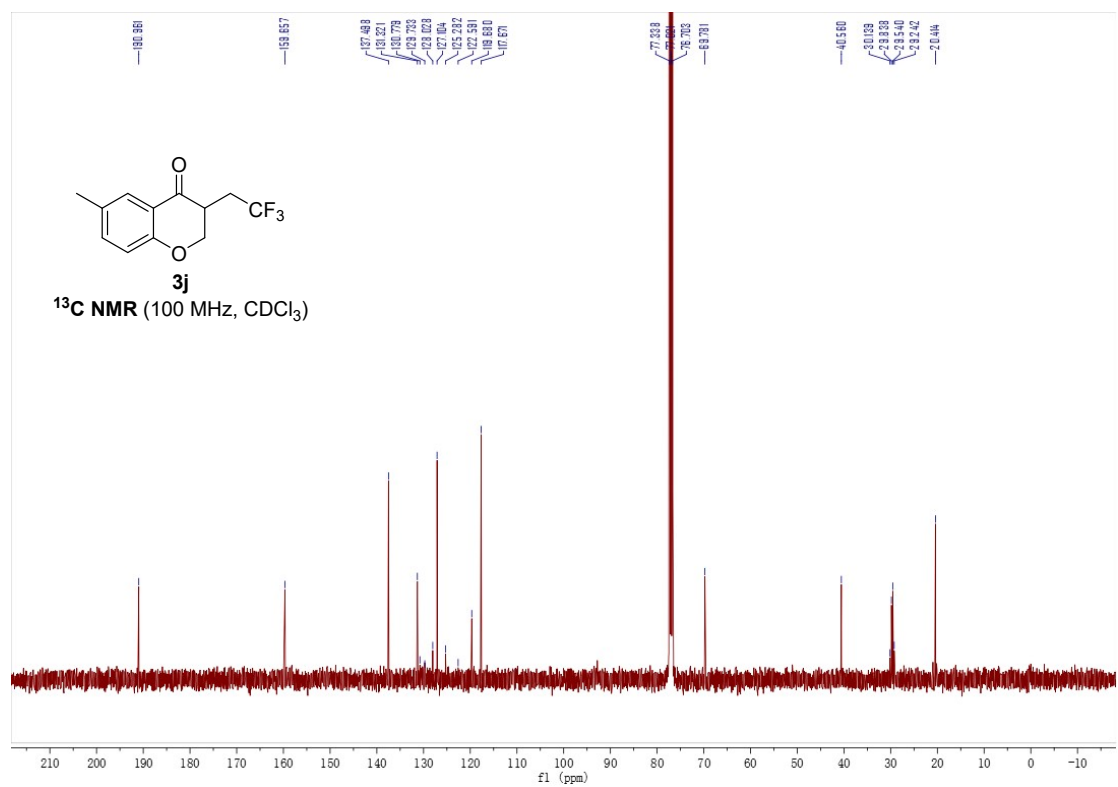
7-nitro-3-(2,2,2-trifluoroethyl)chroman-4-one (3i)



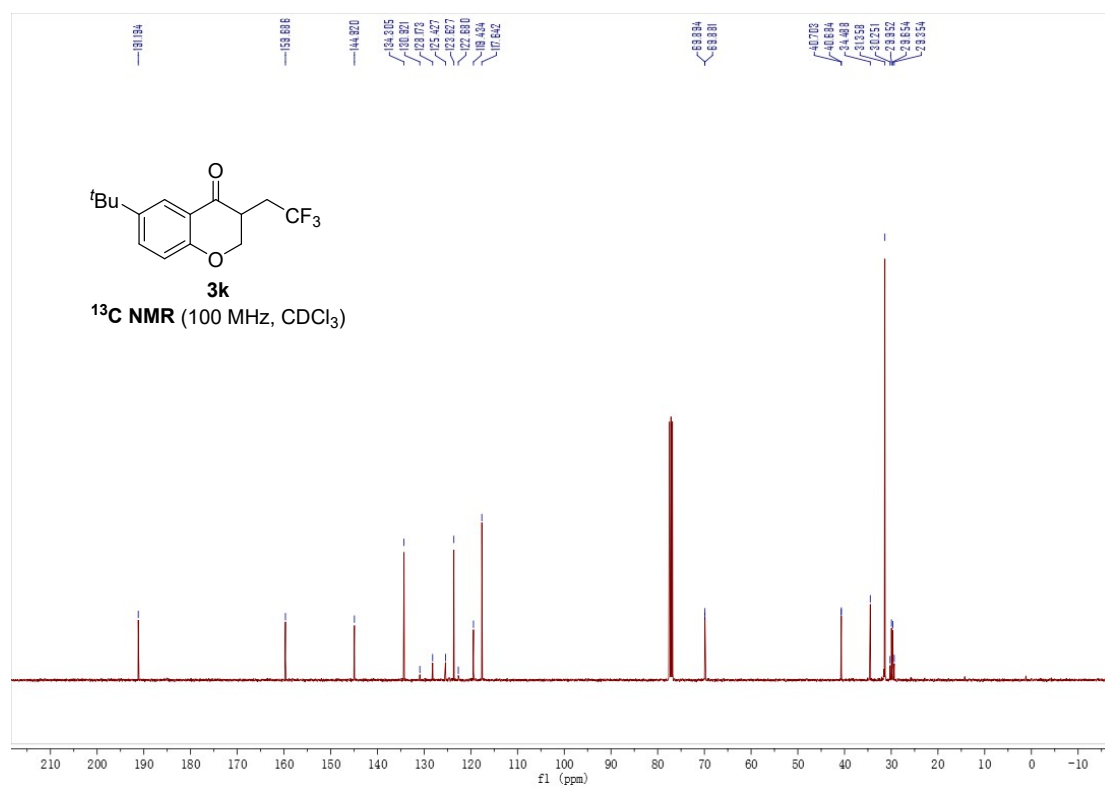
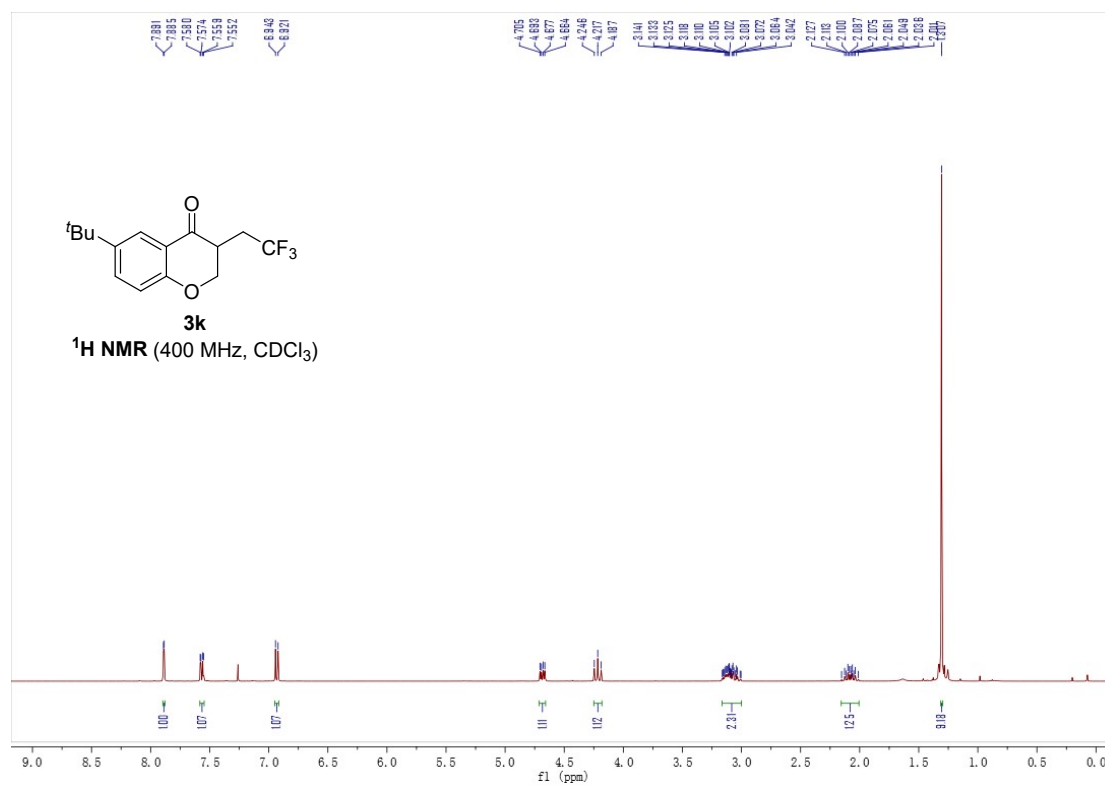


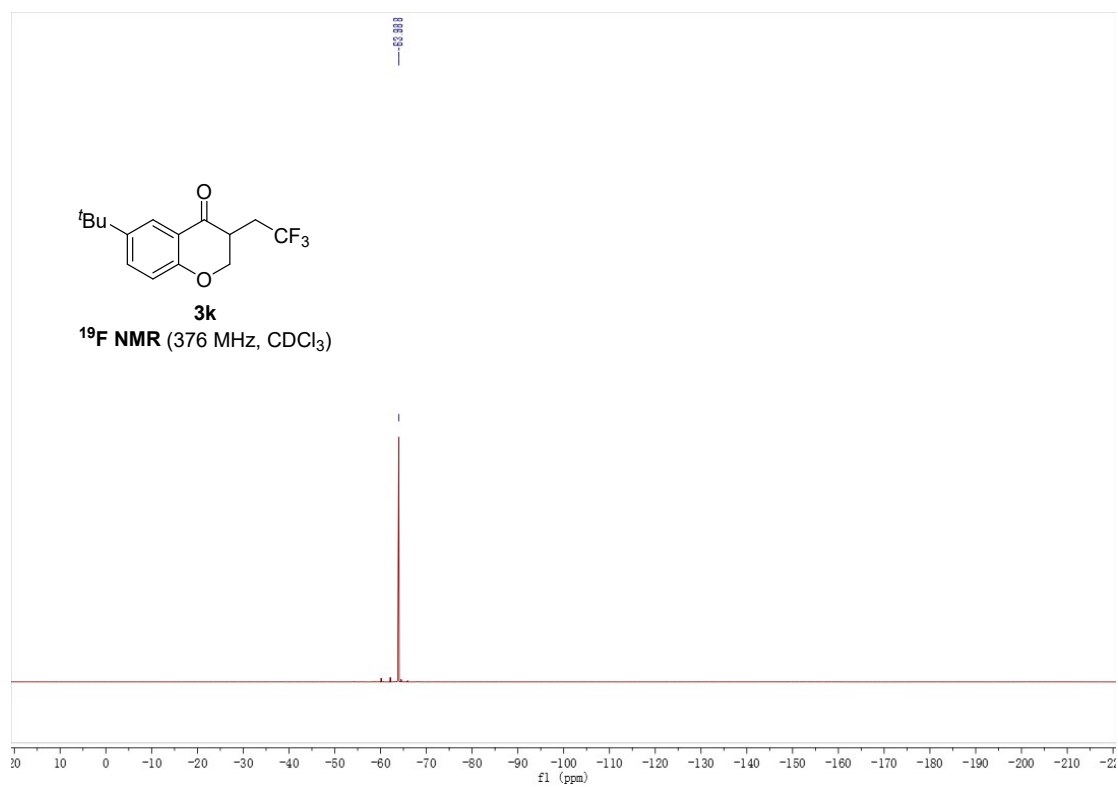
6-methyl-3-(2,2,2-trifluoroethyl)chroman-4-one (3j)



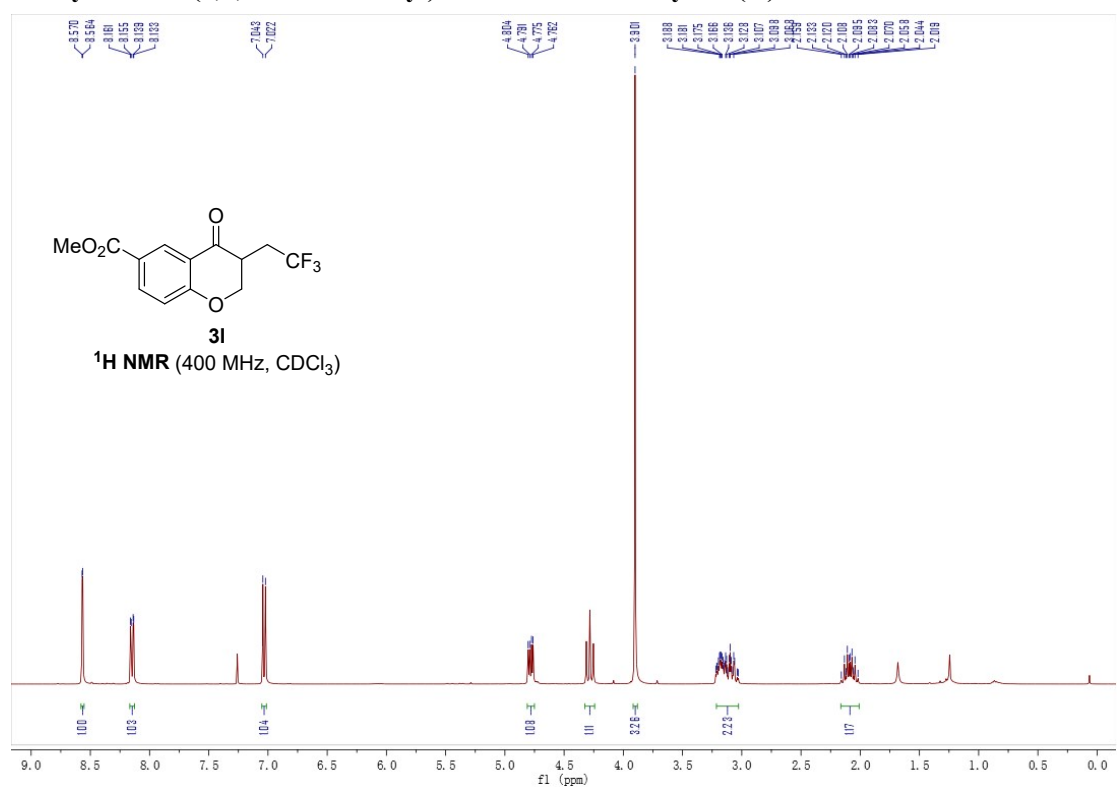


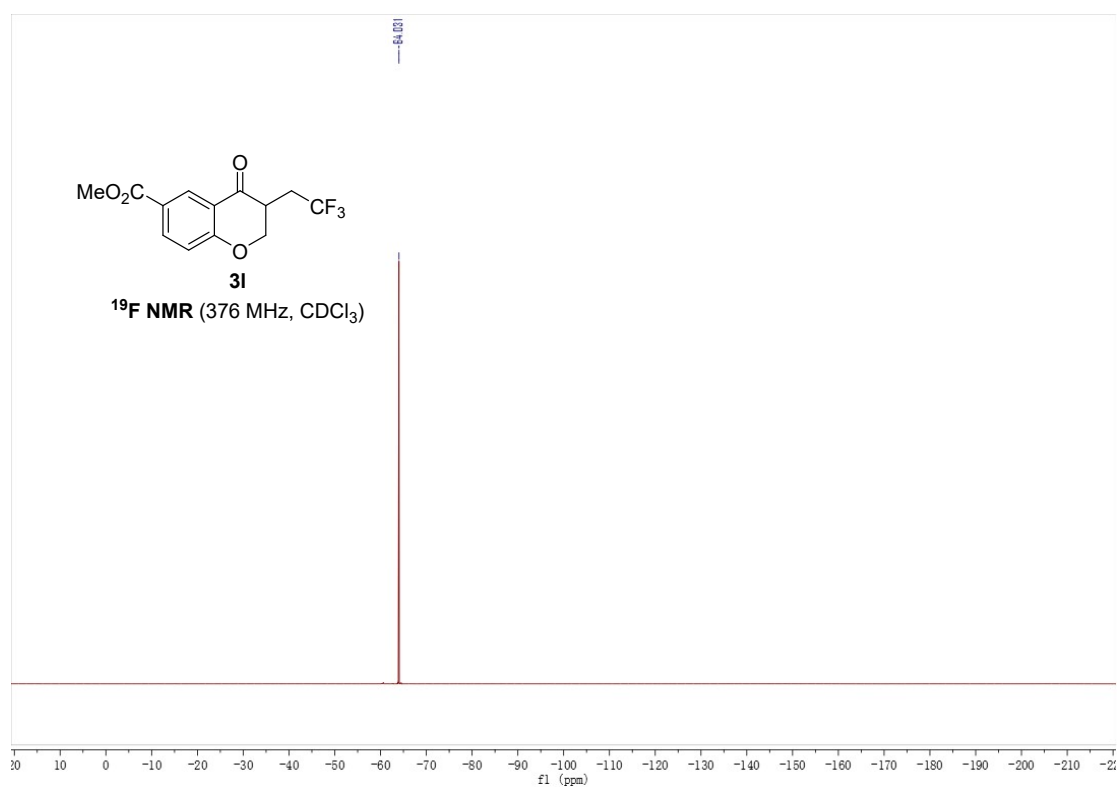
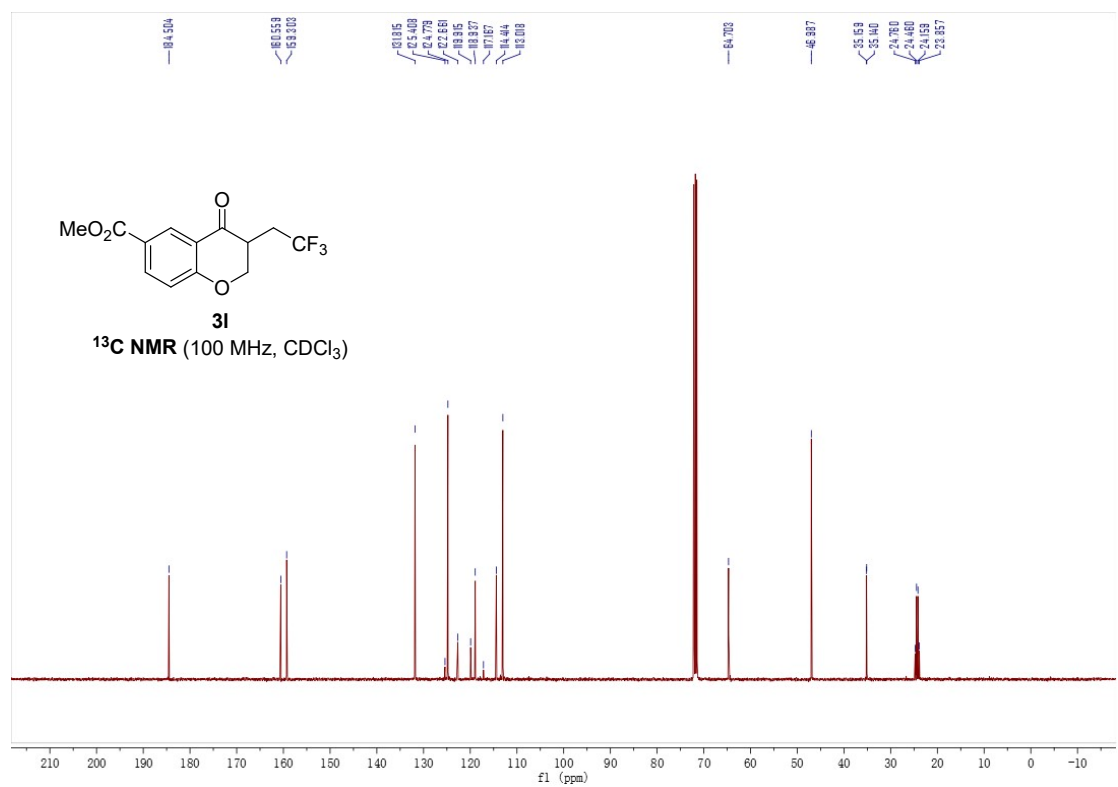
6-(tert-butyl)-3-(2,2,2-trifluoroethyl)chroman-4-one (3k)



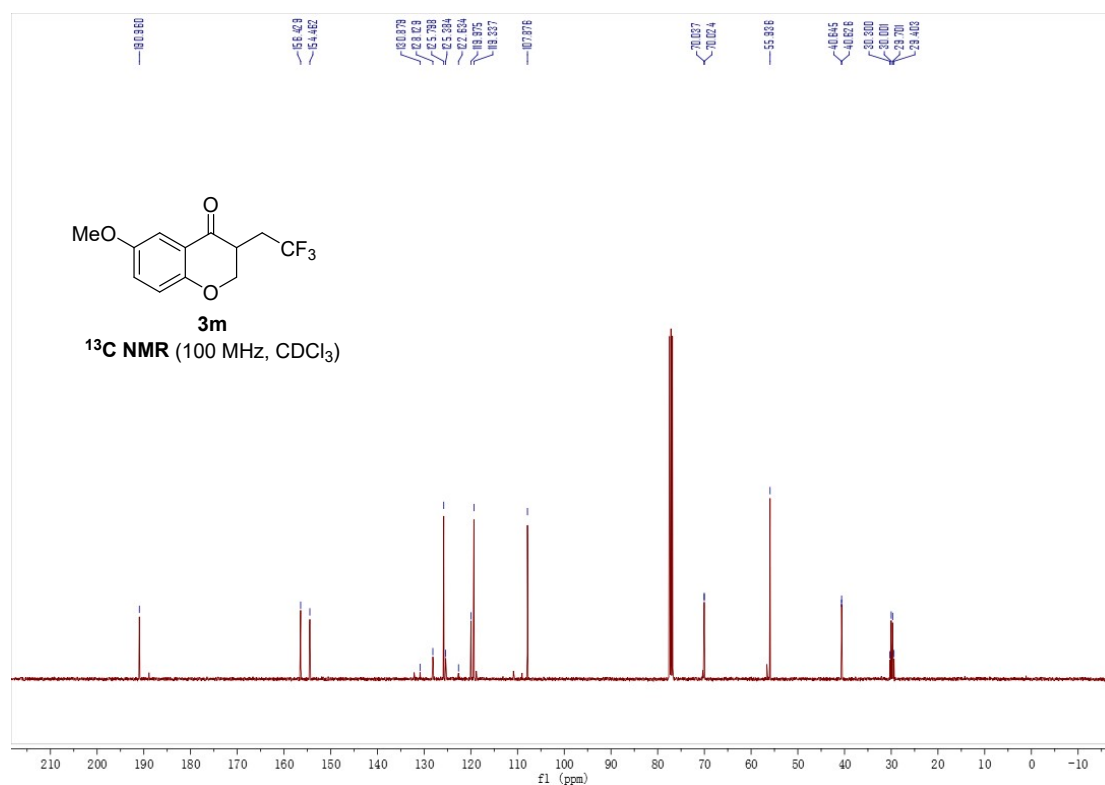
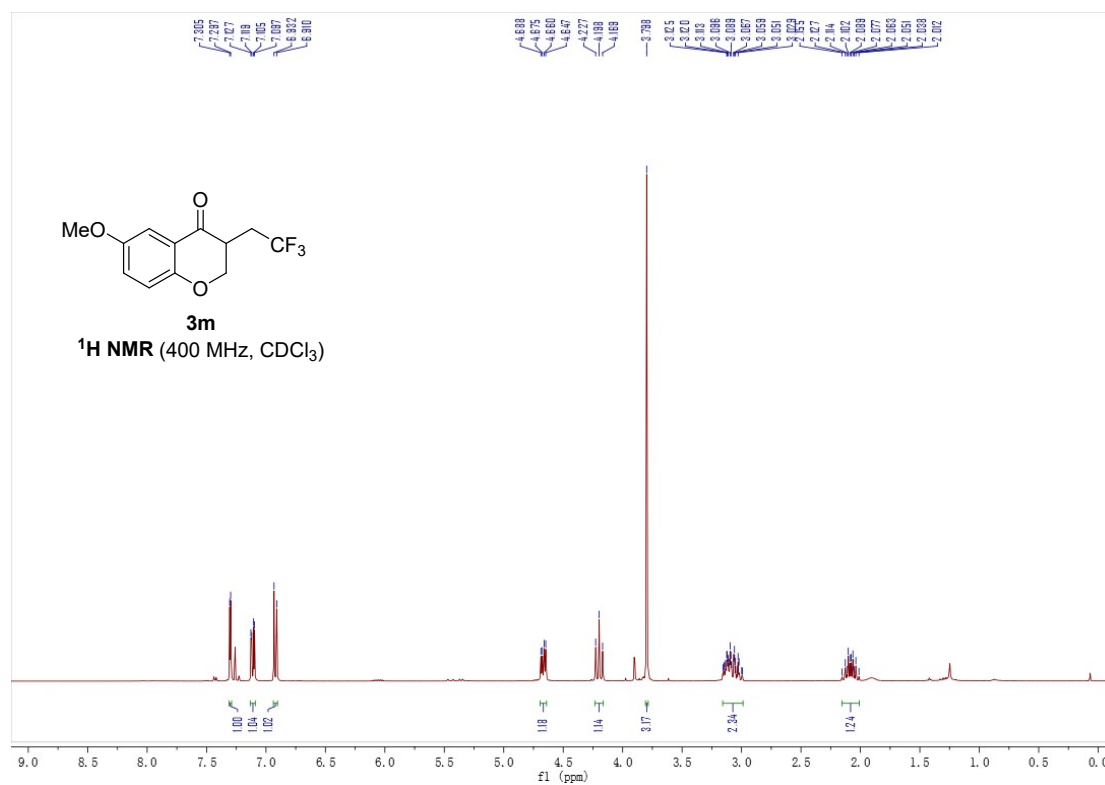


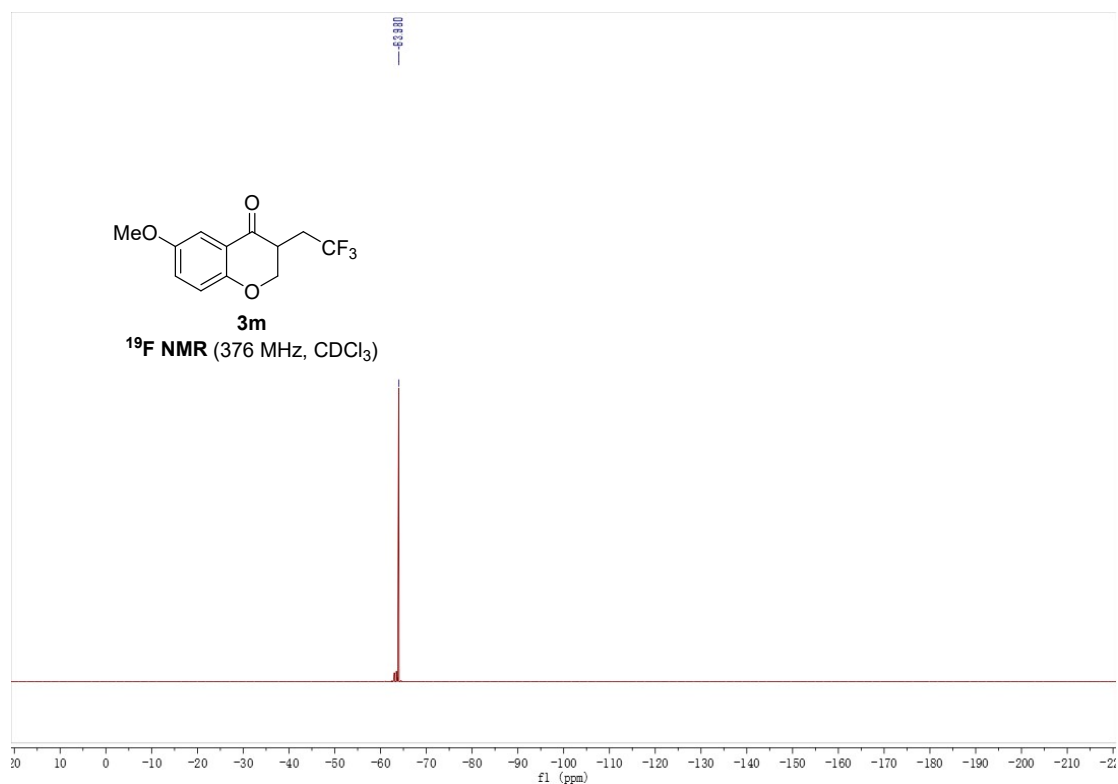
methyl 4-oxo-3-(2,2,2-trifluoroethyl)chromane-6-carboxylate (3l)



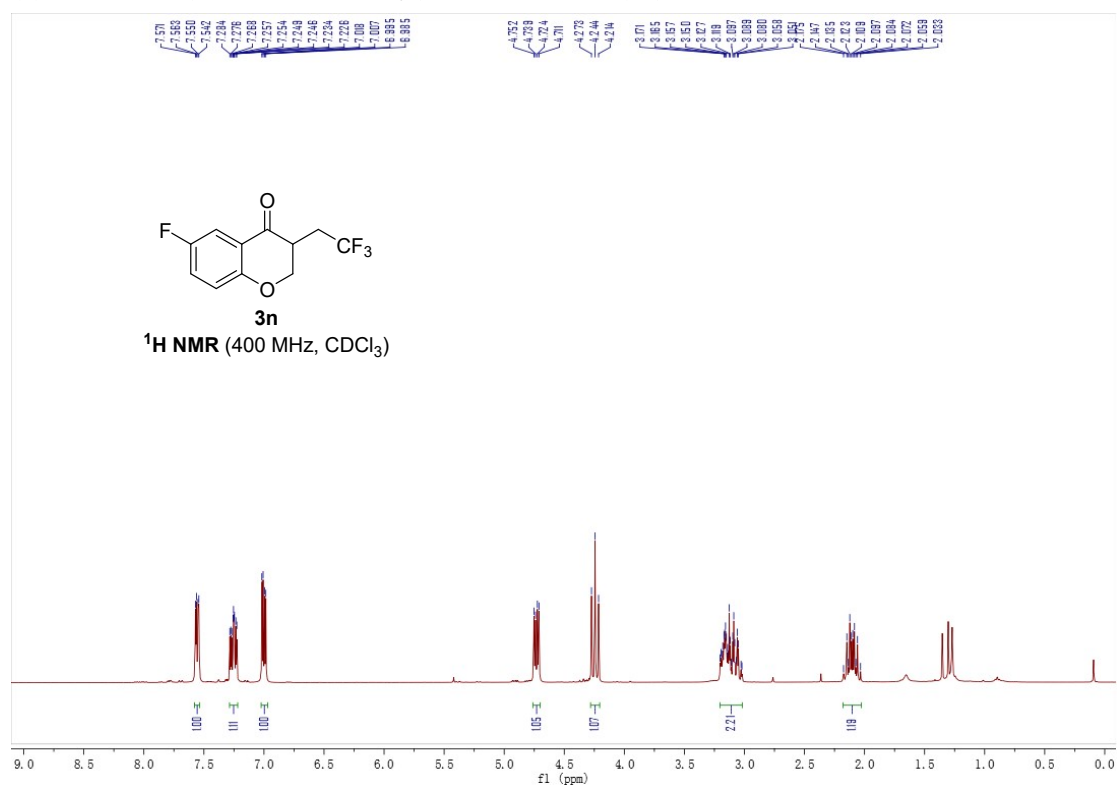


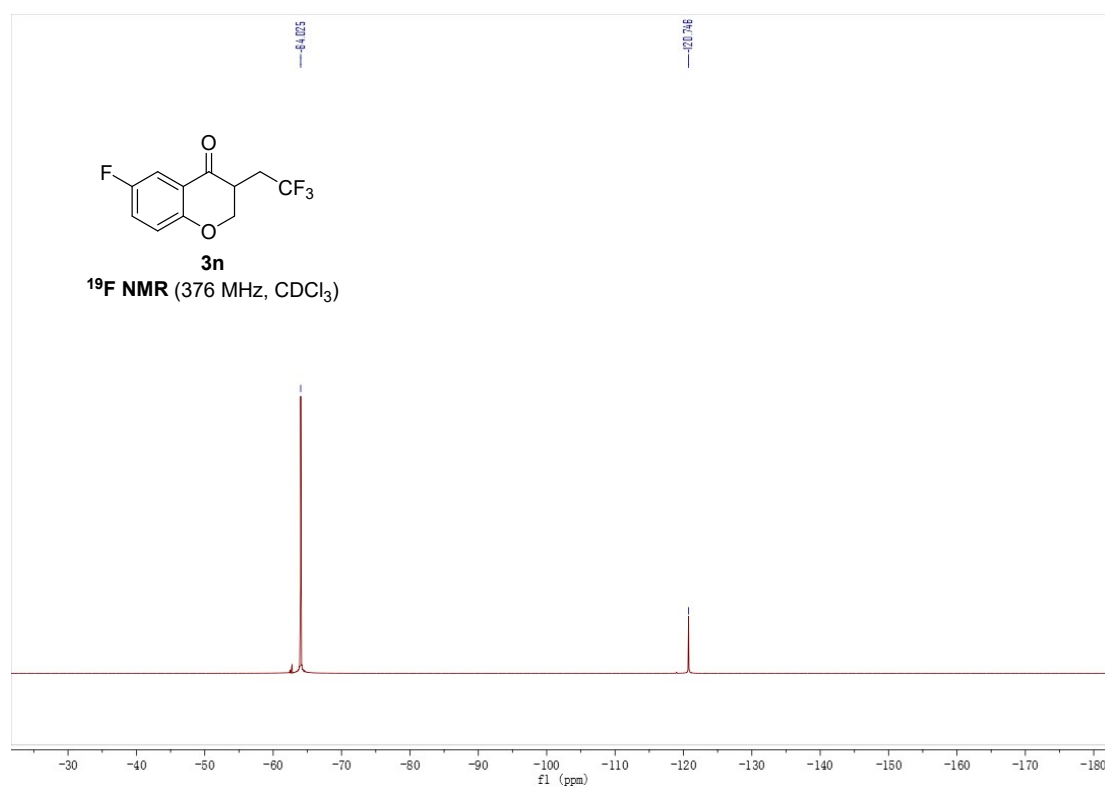
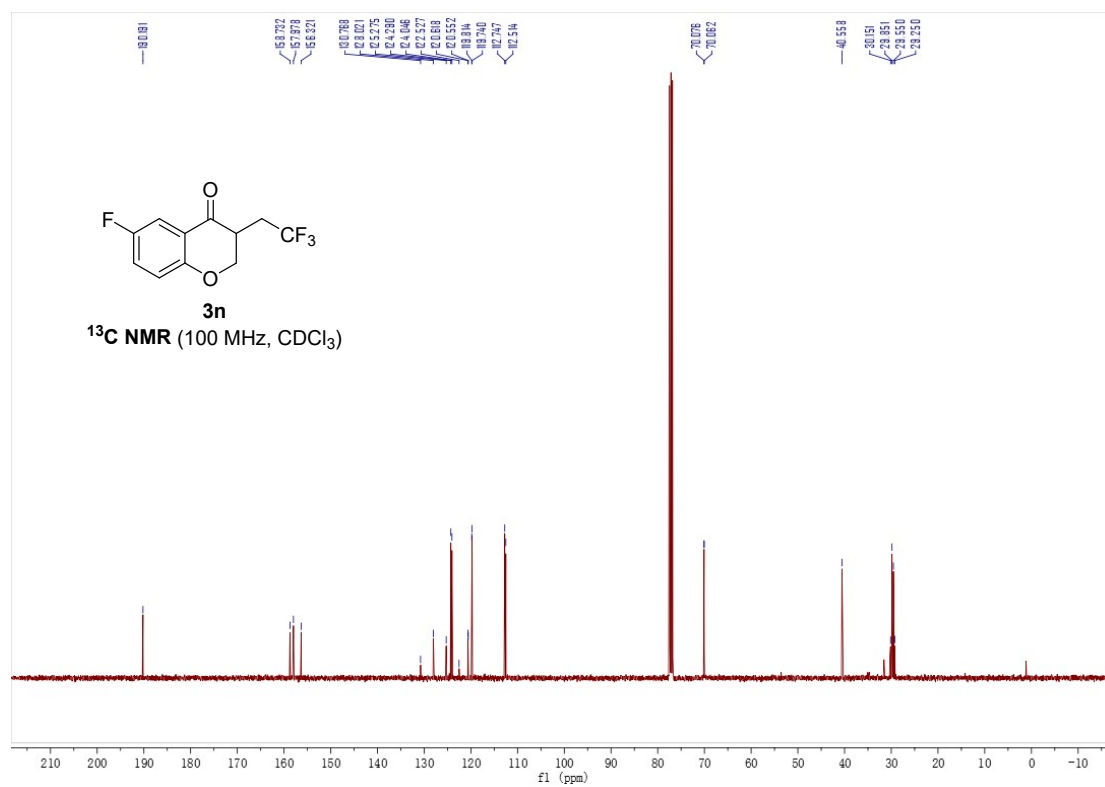
(S)-6-methoxy-3-(2,2,2-trifluoroethyl)chroman-4-one (3m)



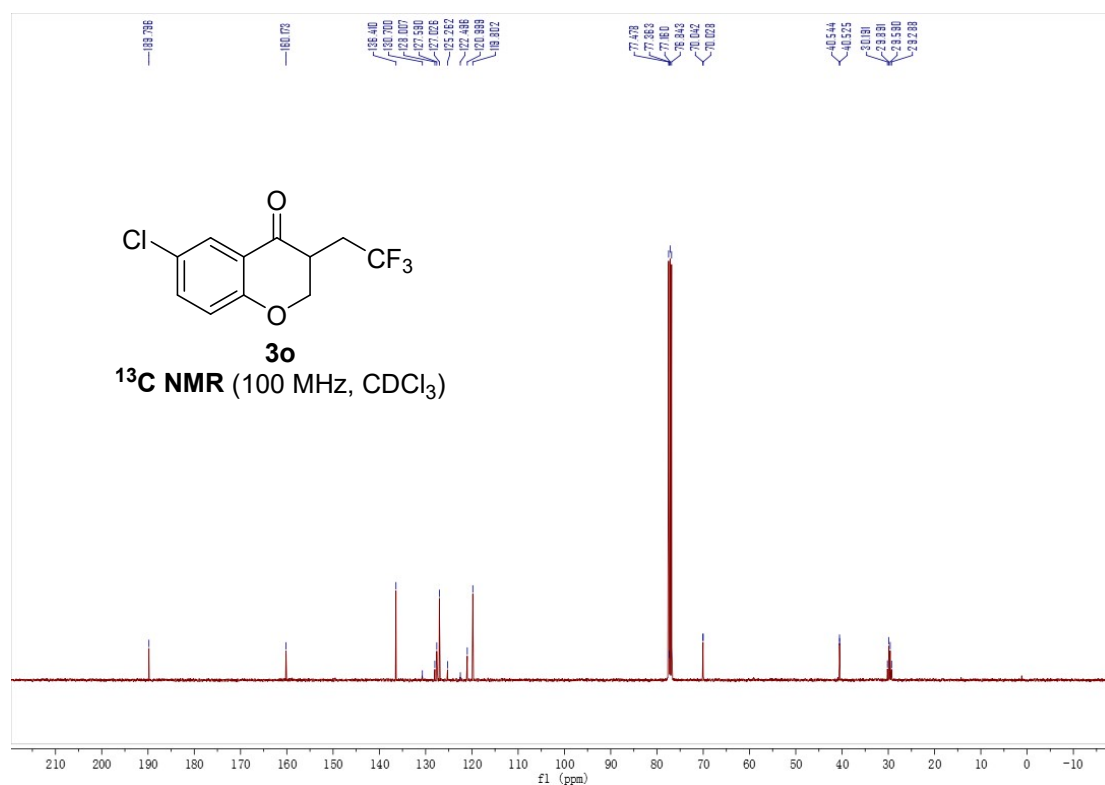
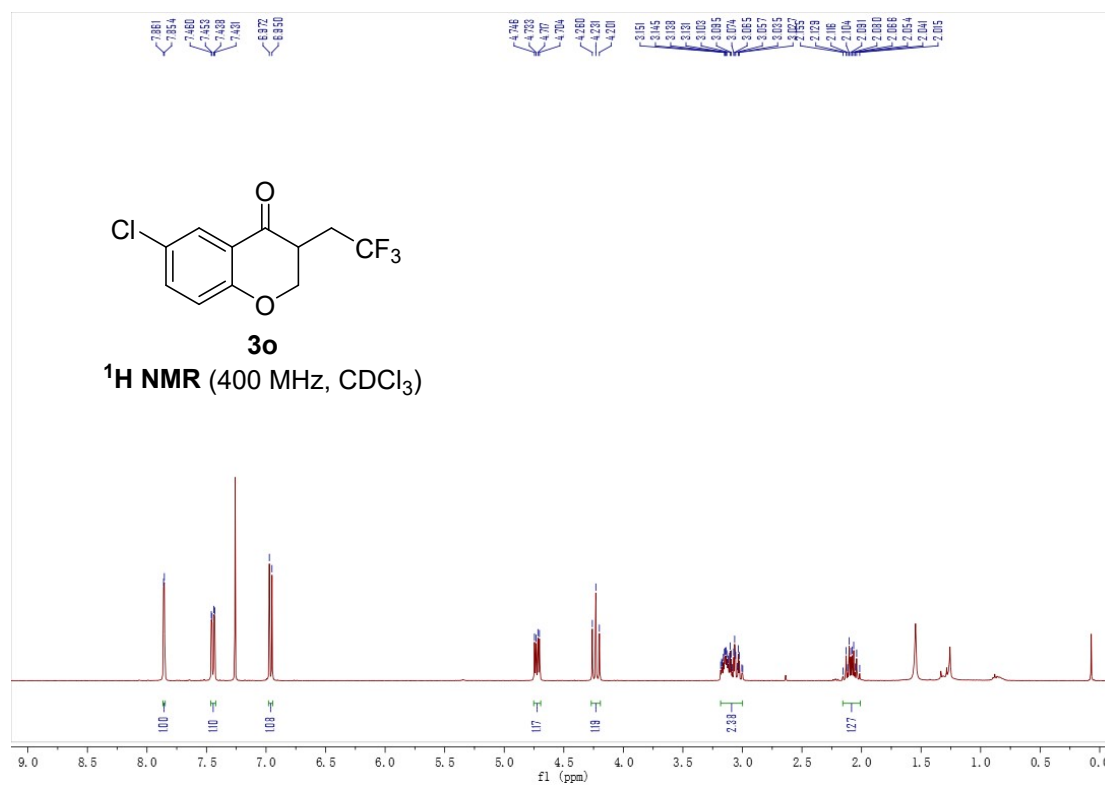


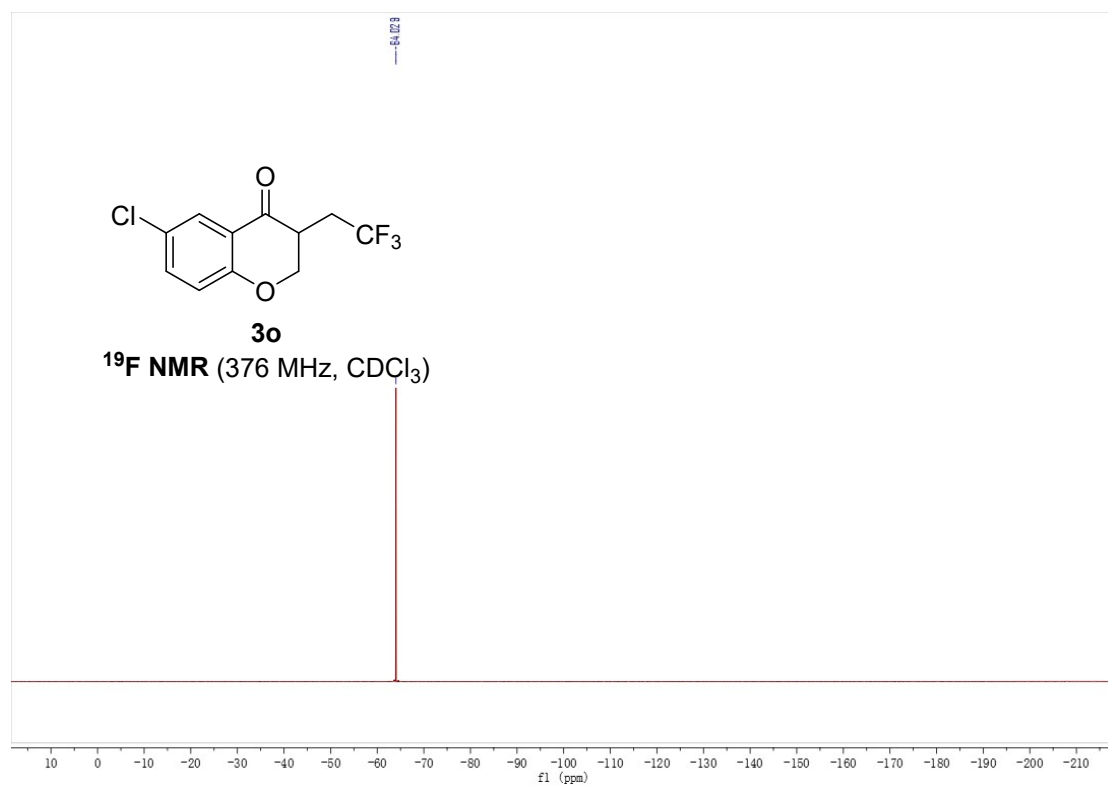
(S)-6-fluoro-3-(2,2,2-trifluoroethyl)chroman-4-one (3n)



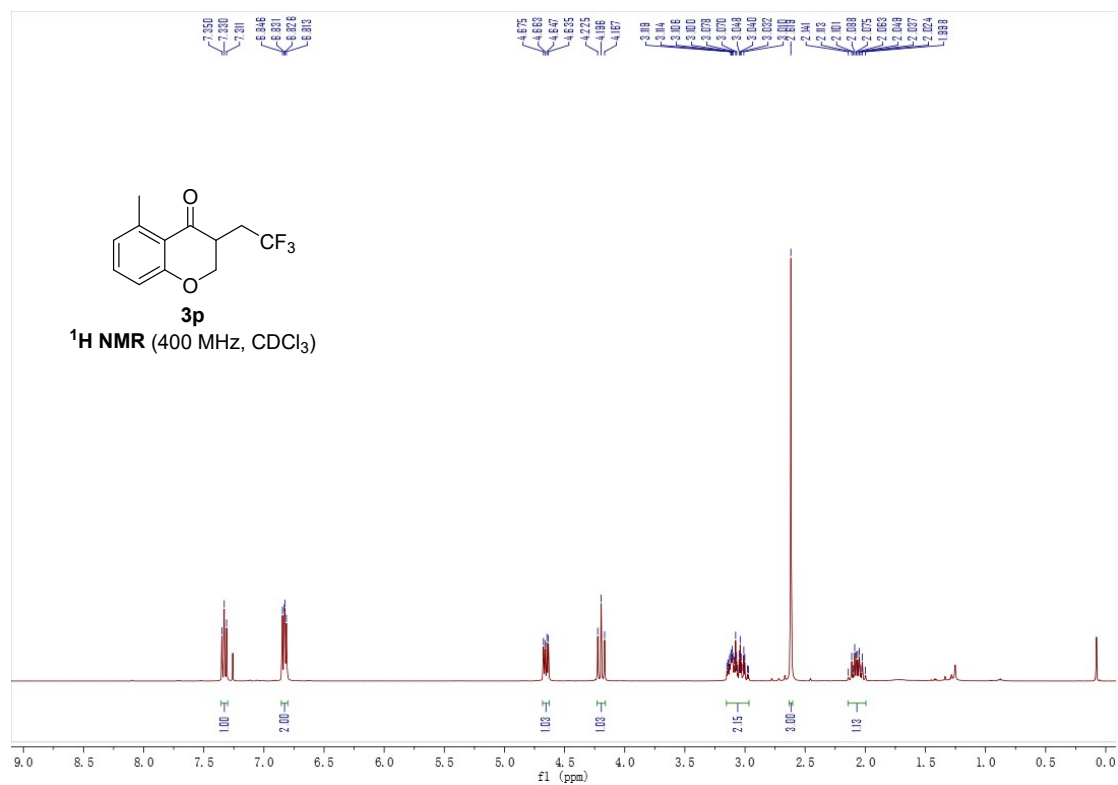


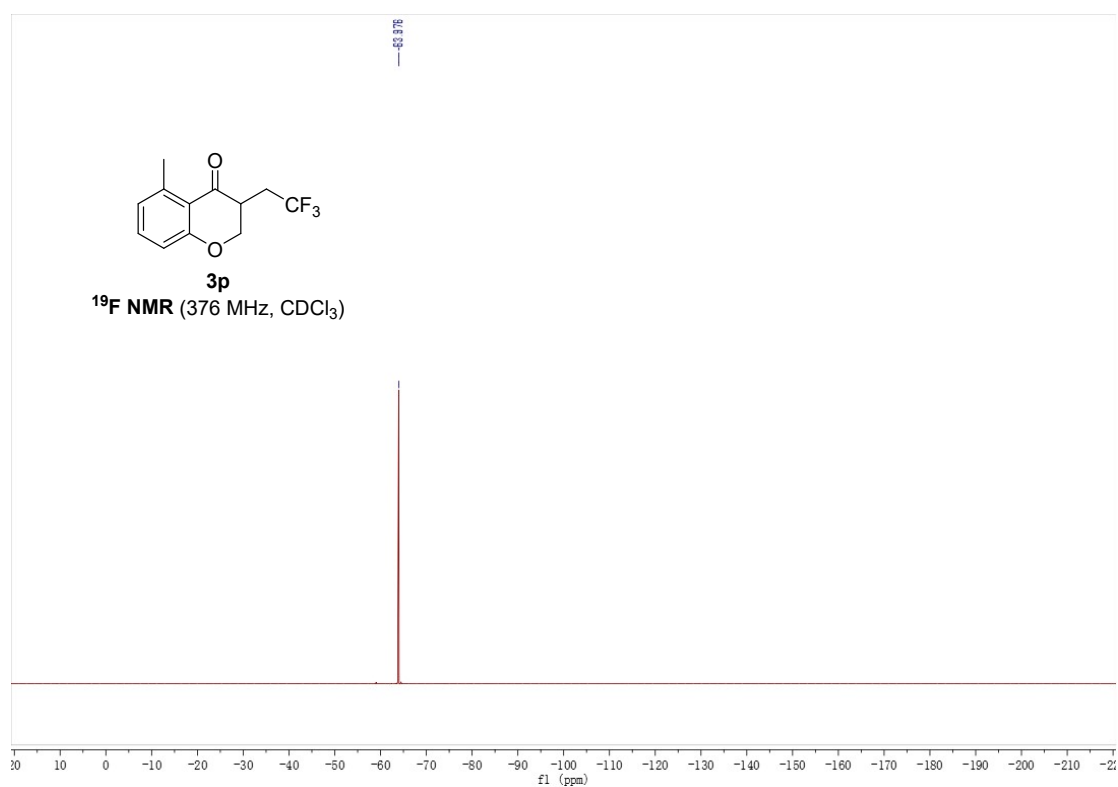
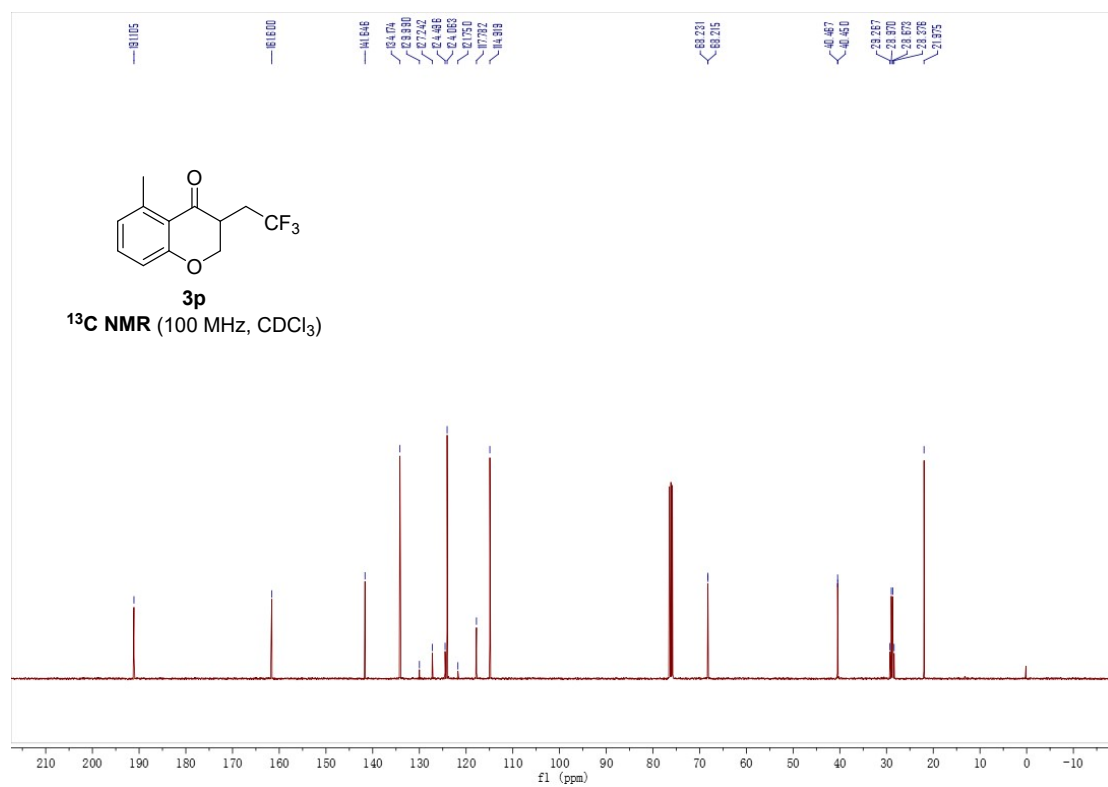
(S)-6-chloro-3-(2,2,2-trifluoroethyl)chroman-4-one (3o)





5-methyl-3-(2,2,2-trifluoroethyl)chroman-4-one (3p)

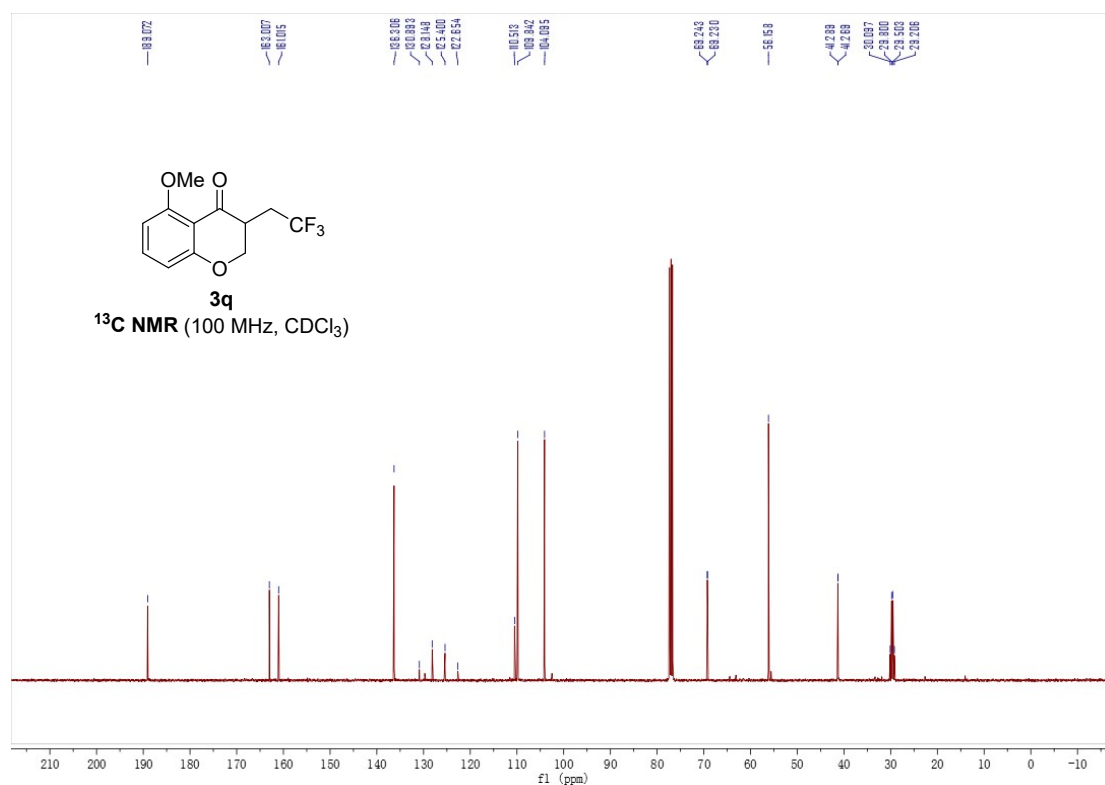


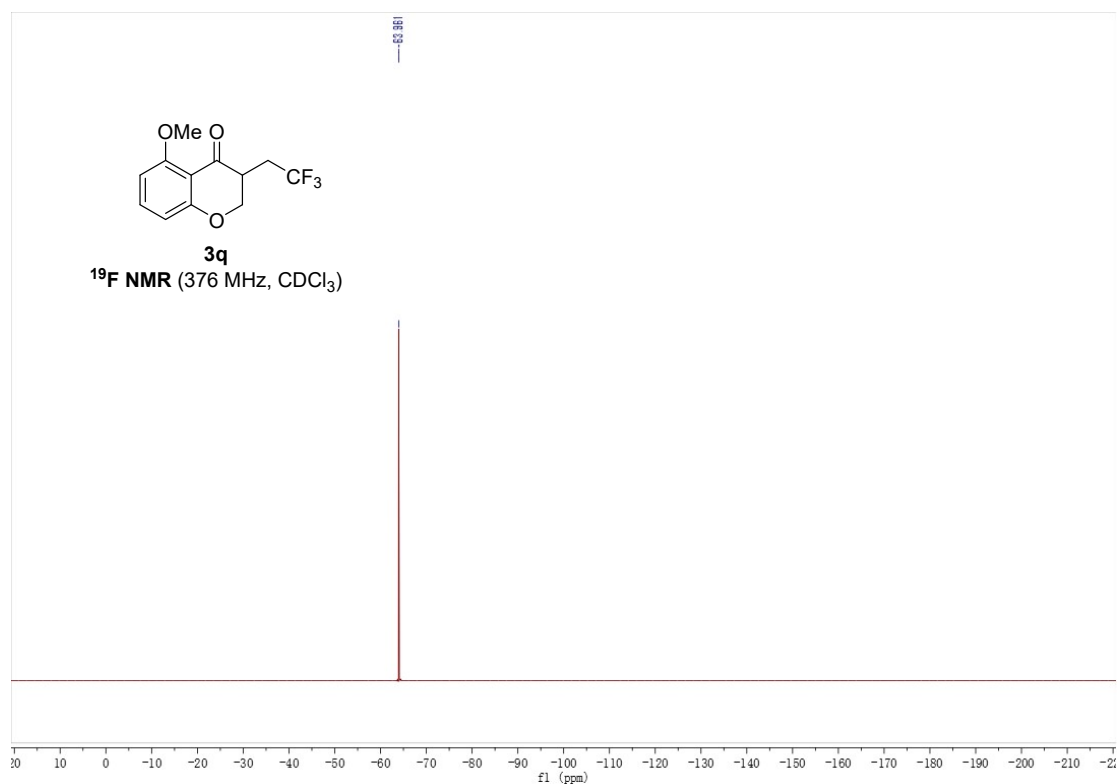


3q
¹H NMR (400 MHz, CDCl₃)

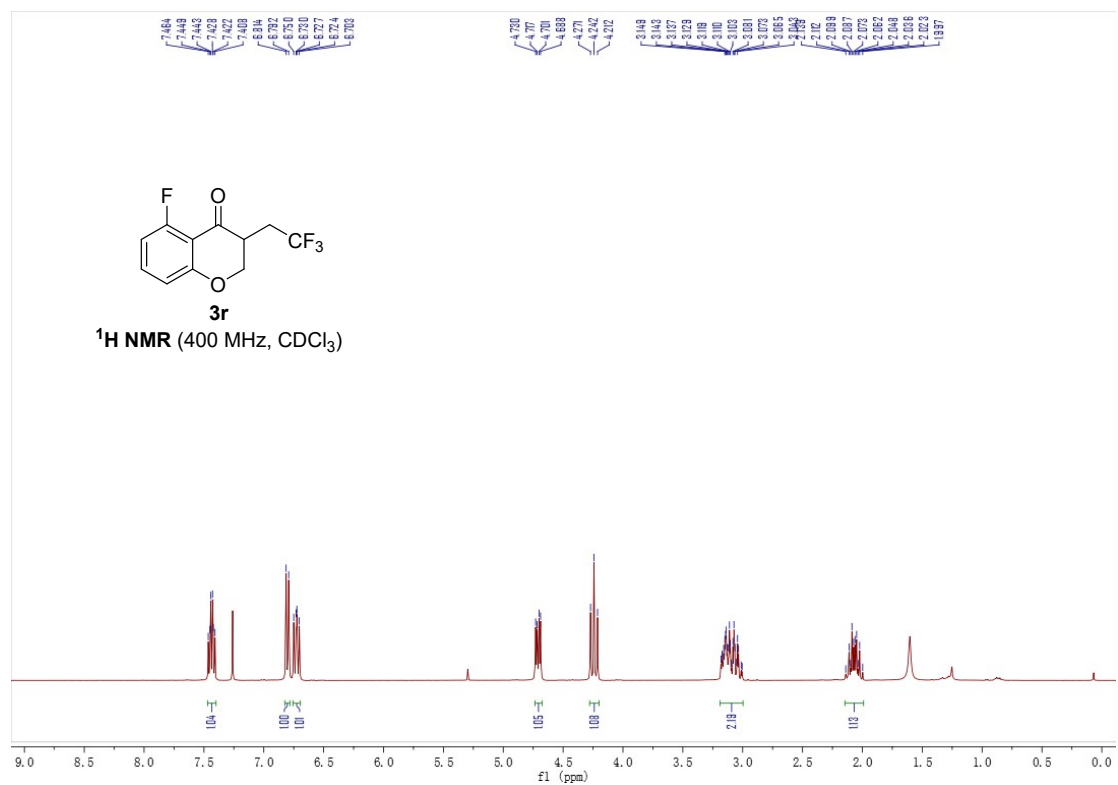
Chemical structure of **3q**: COC1=CC=C2C(=C1)OC(CC2=O)COC

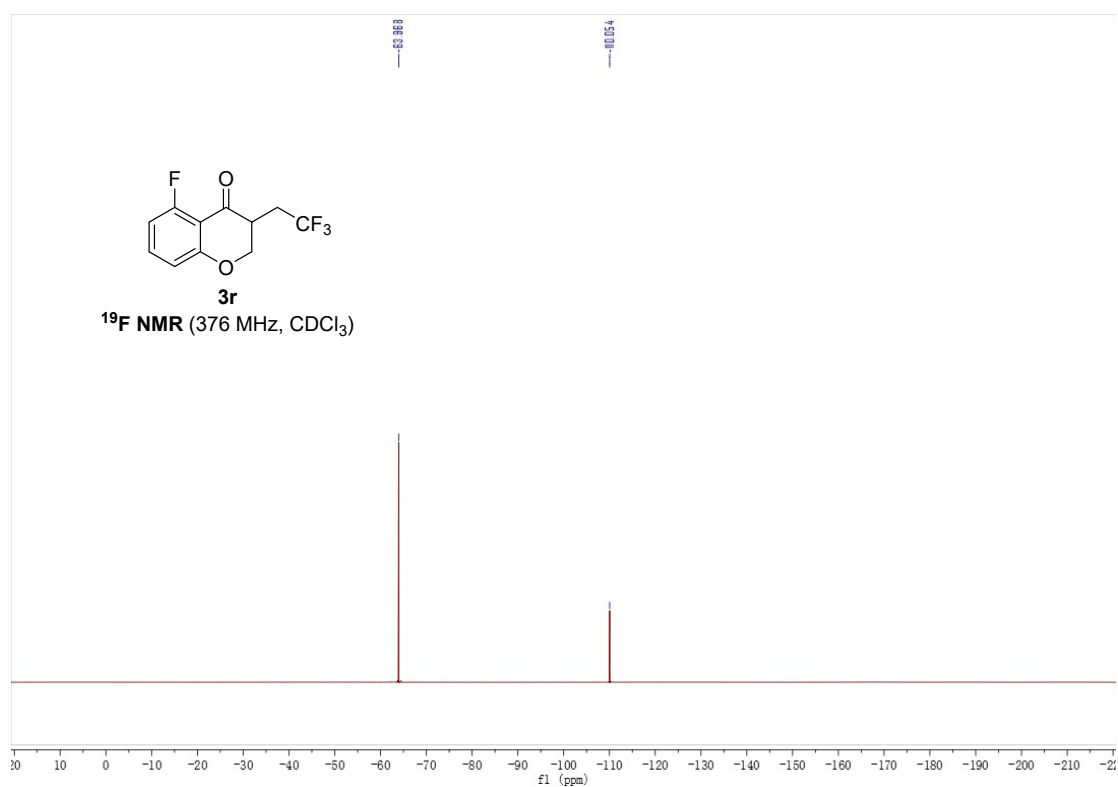
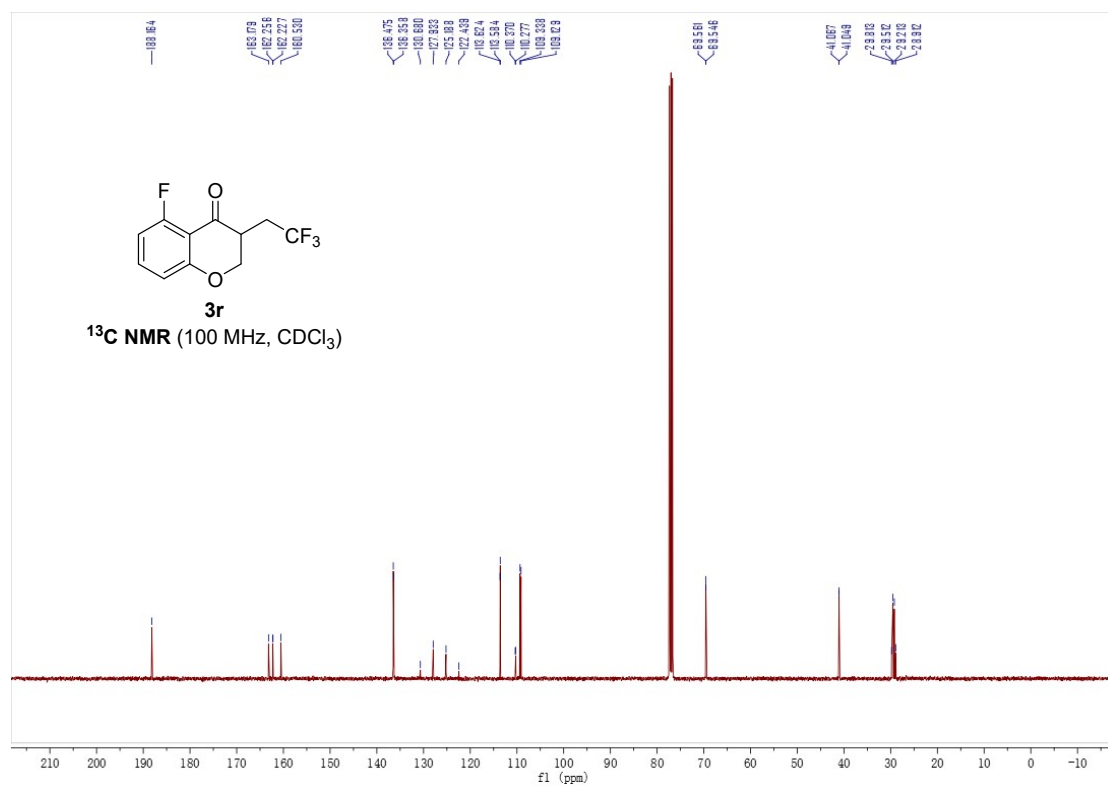
¹H NMR spectrum (400 MHz, CDCl₃) showing peaks from 0.0 to 9.0 ppm. The spectrum includes a sharp singlet at ~3.8 ppm (3H, OMe), a multiplet at ~3.1 ppm (2H, CH₂), a multiplet at ~2.1 ppm (2H, CH₂), a multiplet at ~6.5 ppm (2H, aromatic), and a multiplet at ~7.4 ppm (1H, aromatic). Integration values are provided below the baseline.



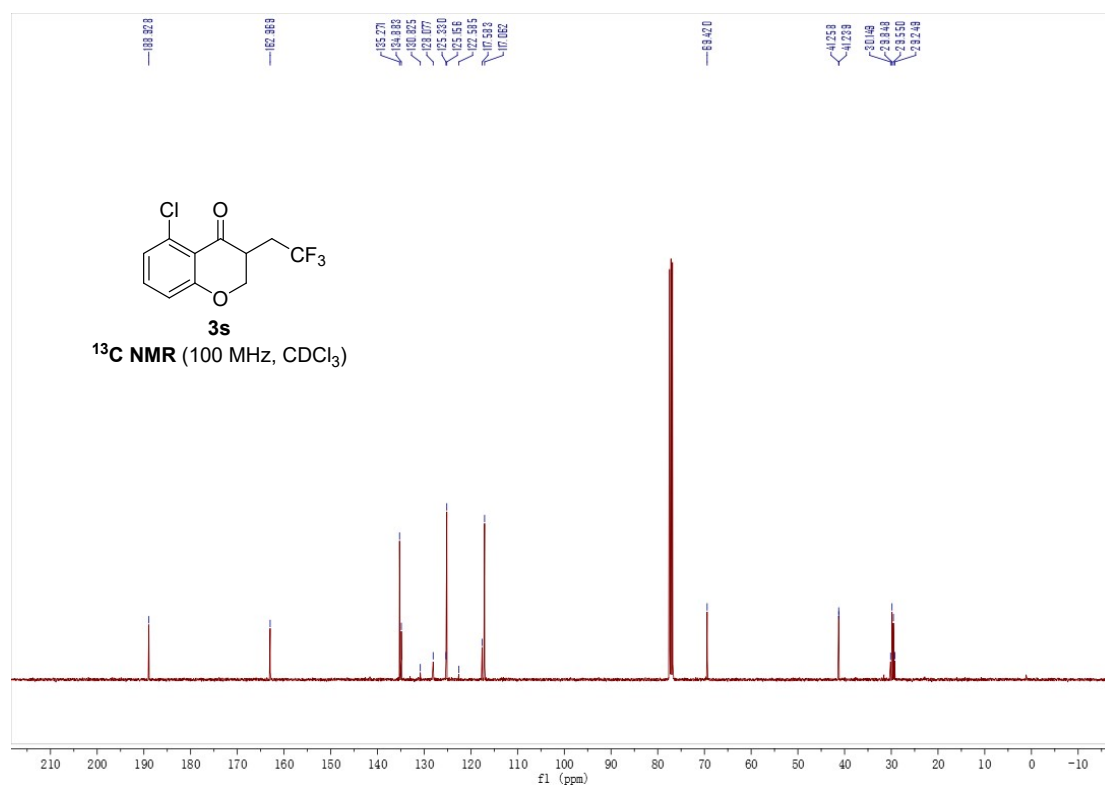
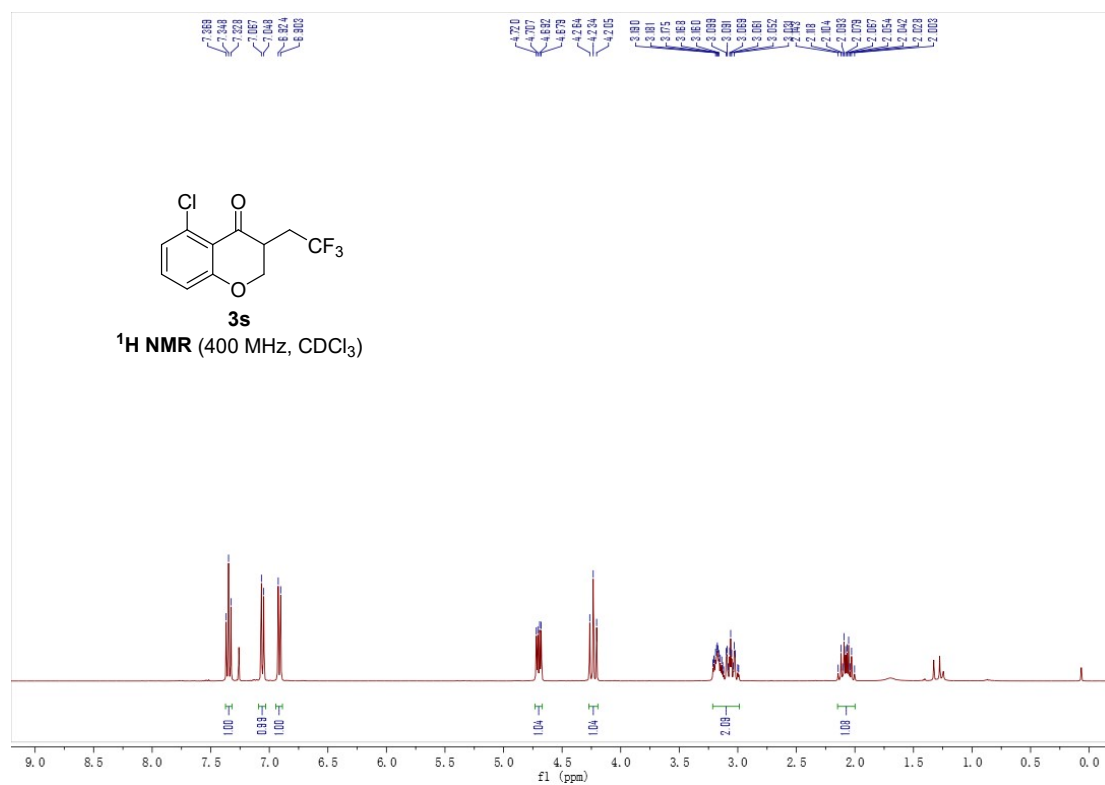


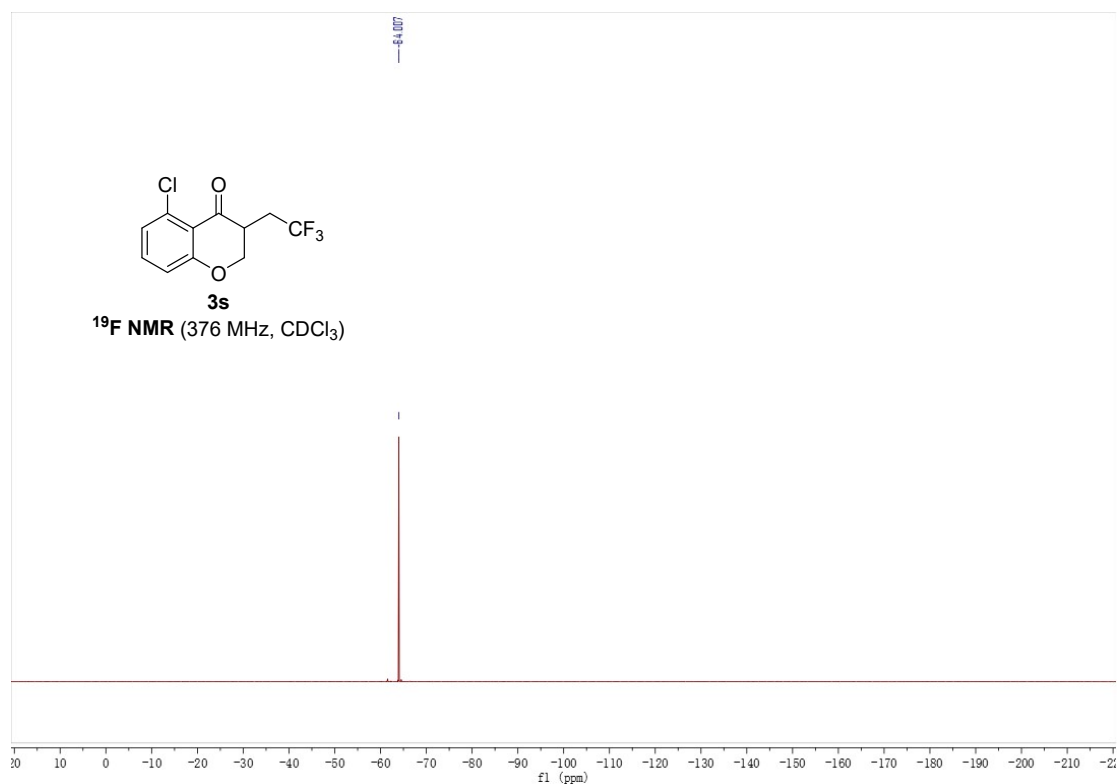
5-fluoro-3-(2,2,2-trifluoroethyl)chroman-4-one (3r)



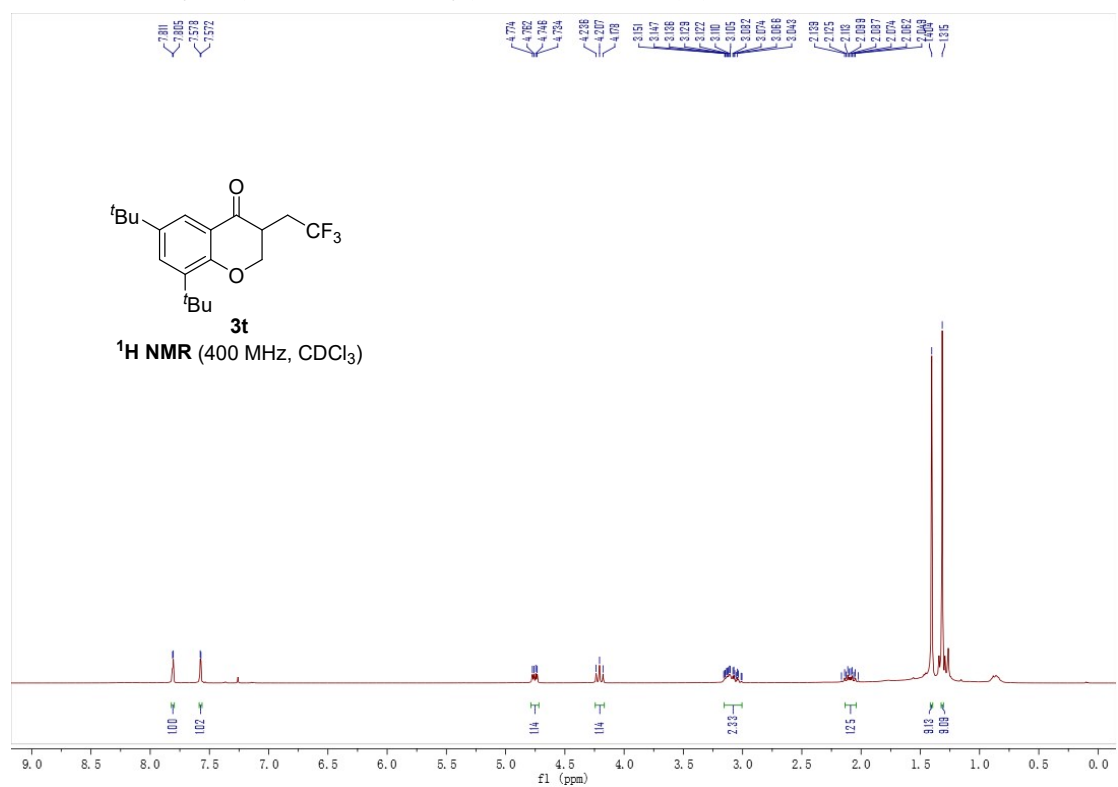


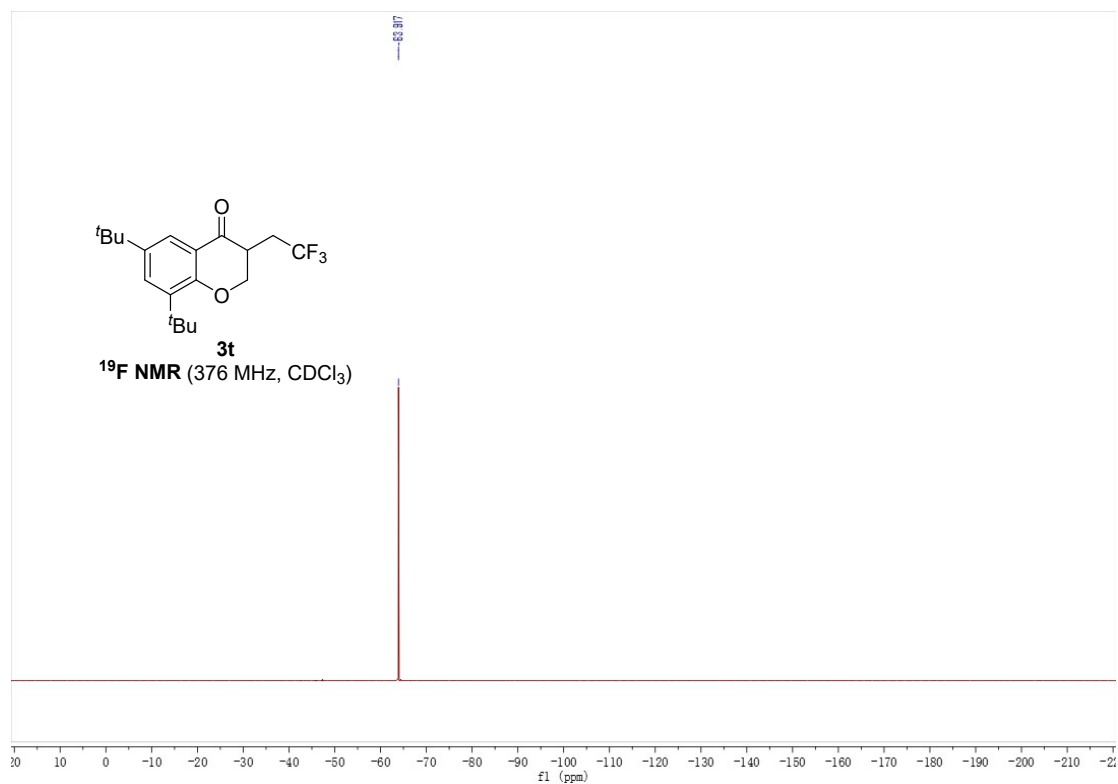
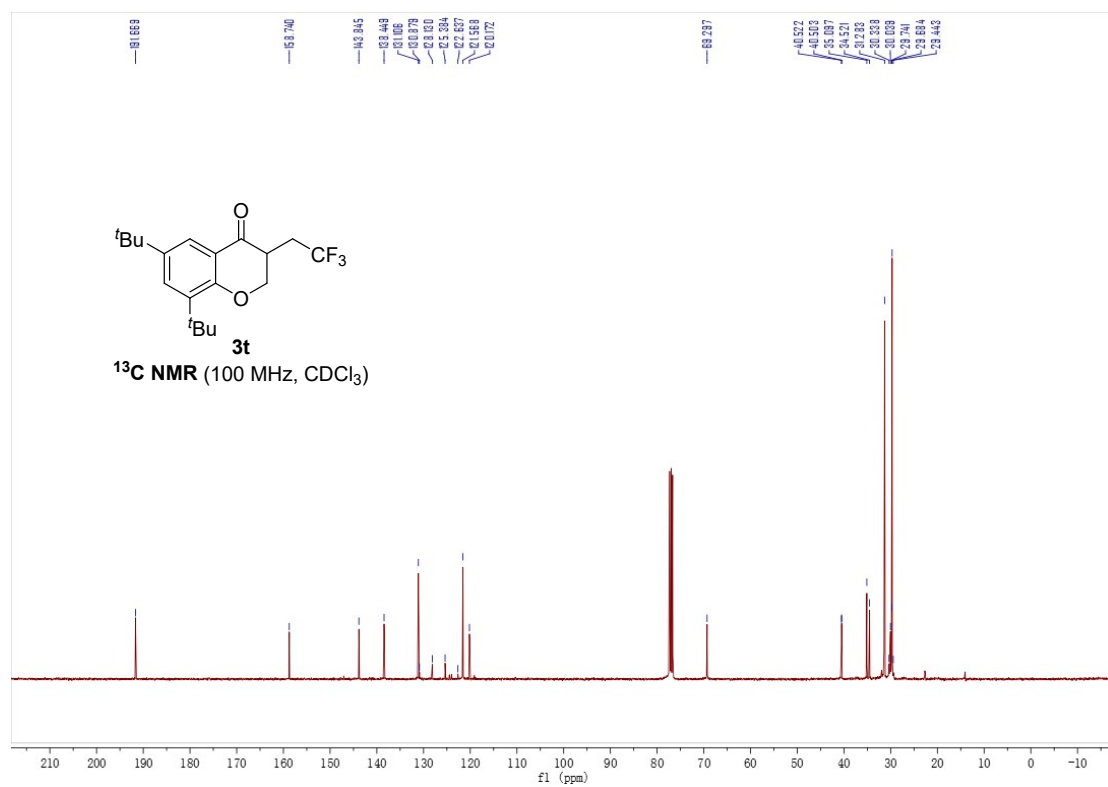
5-chloro-3-(2,2,2-trifluoroethyl)chroman-4-one (3s)



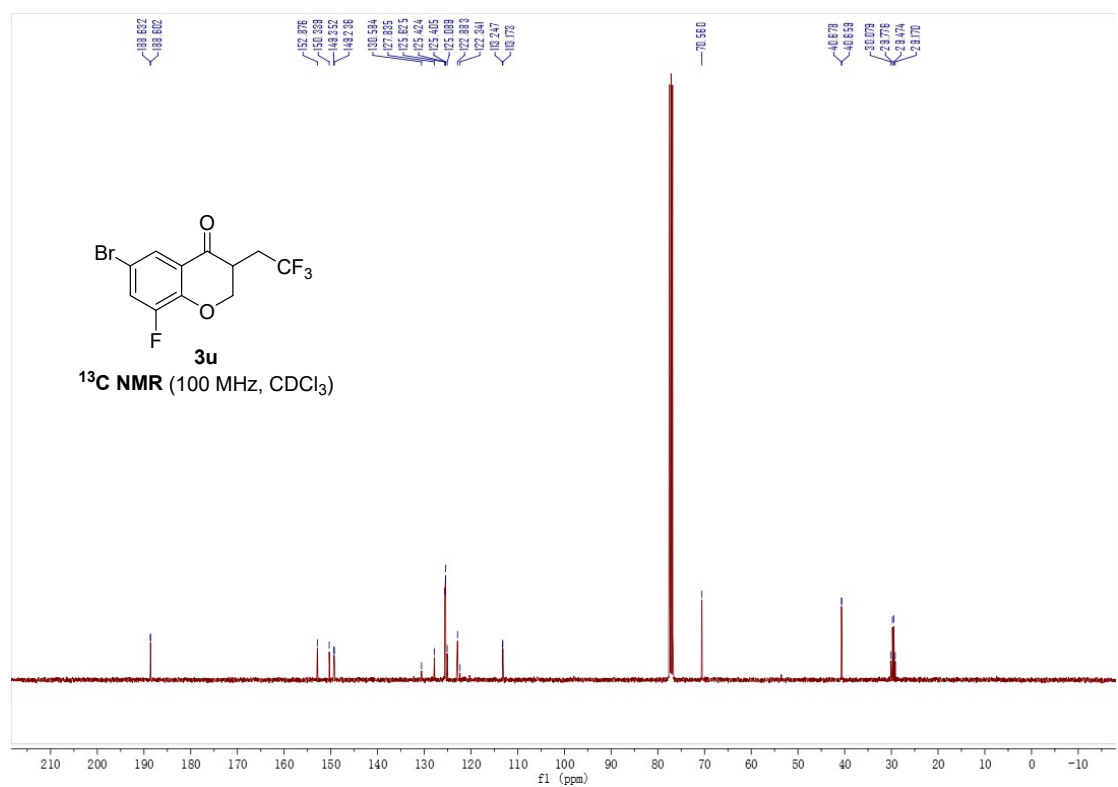
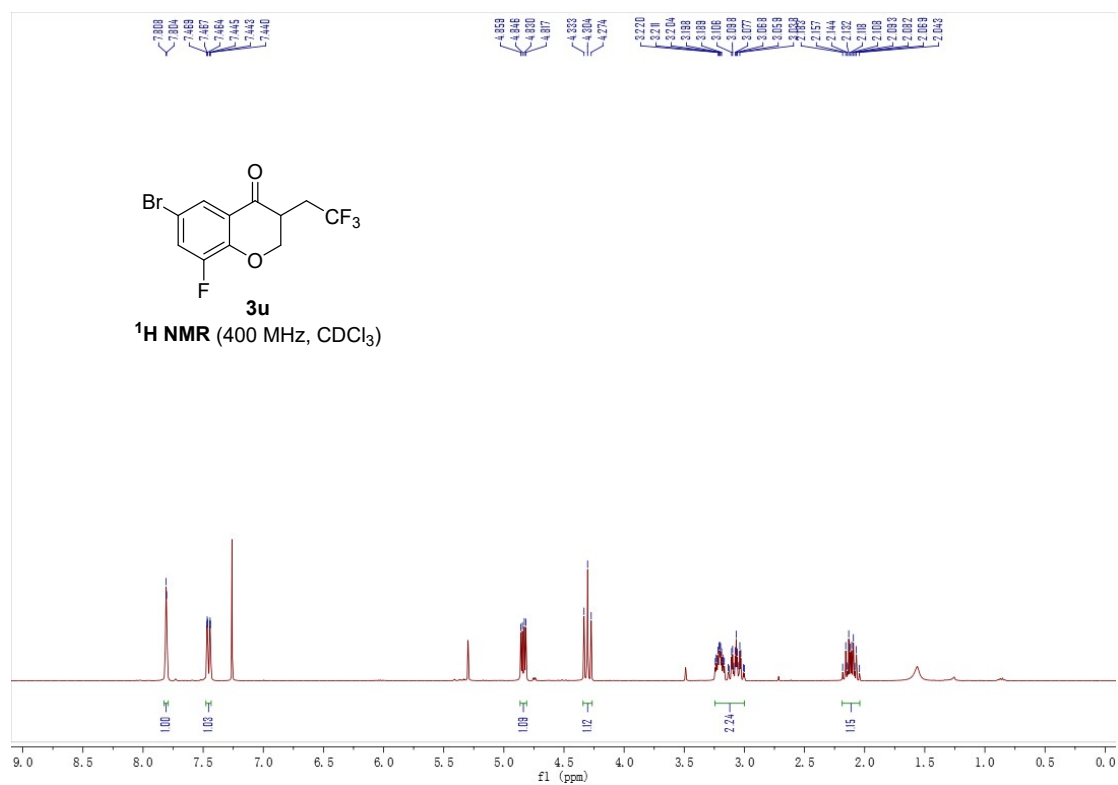


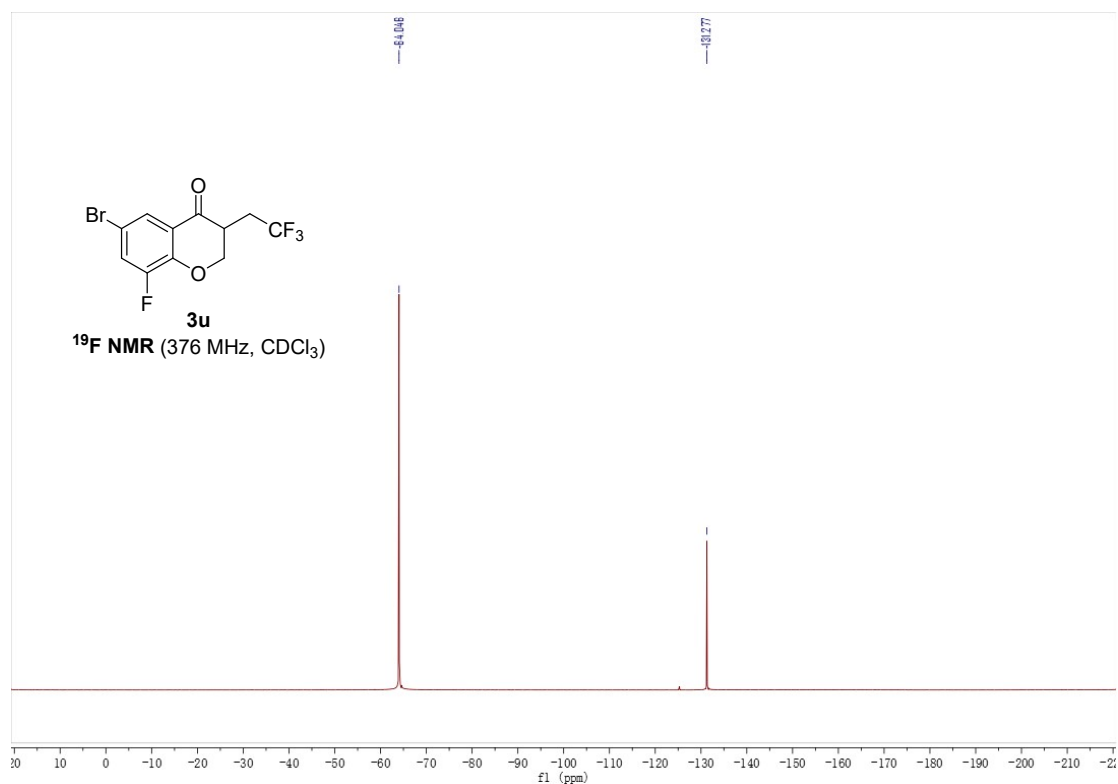
6,8-di-tert-butyl-3-(2,2,2-trifluoroethyl)chroman-4-one (3t)



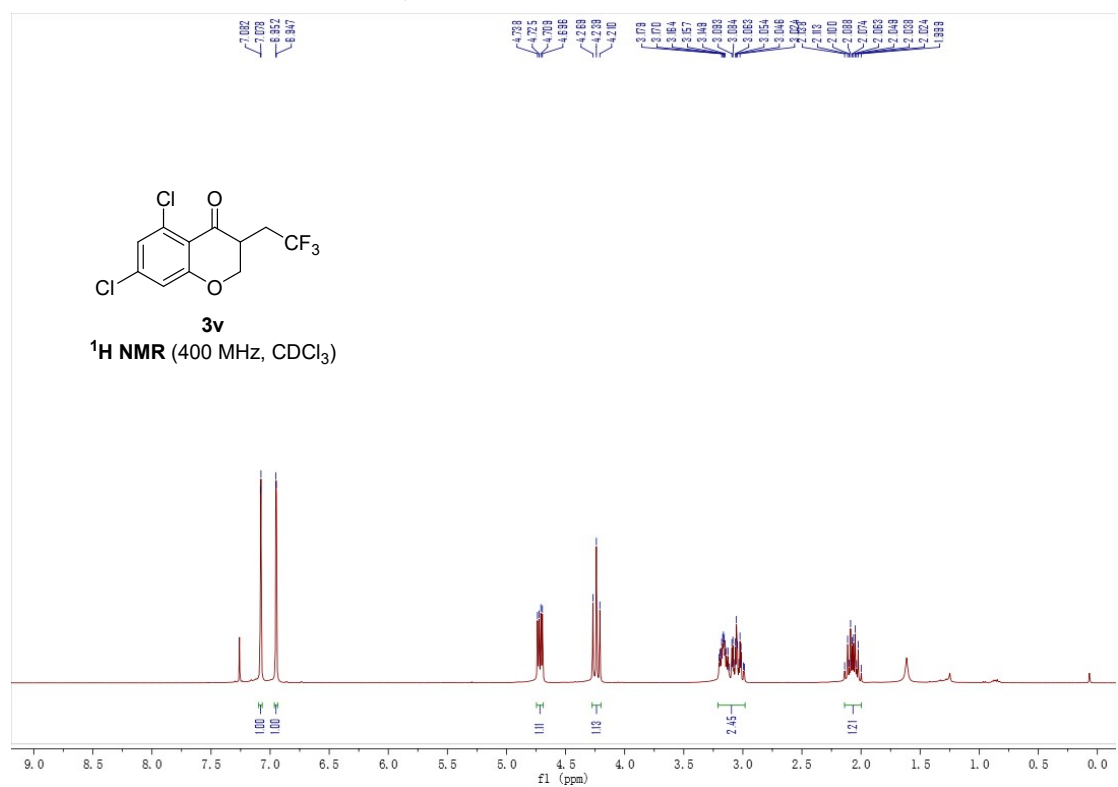


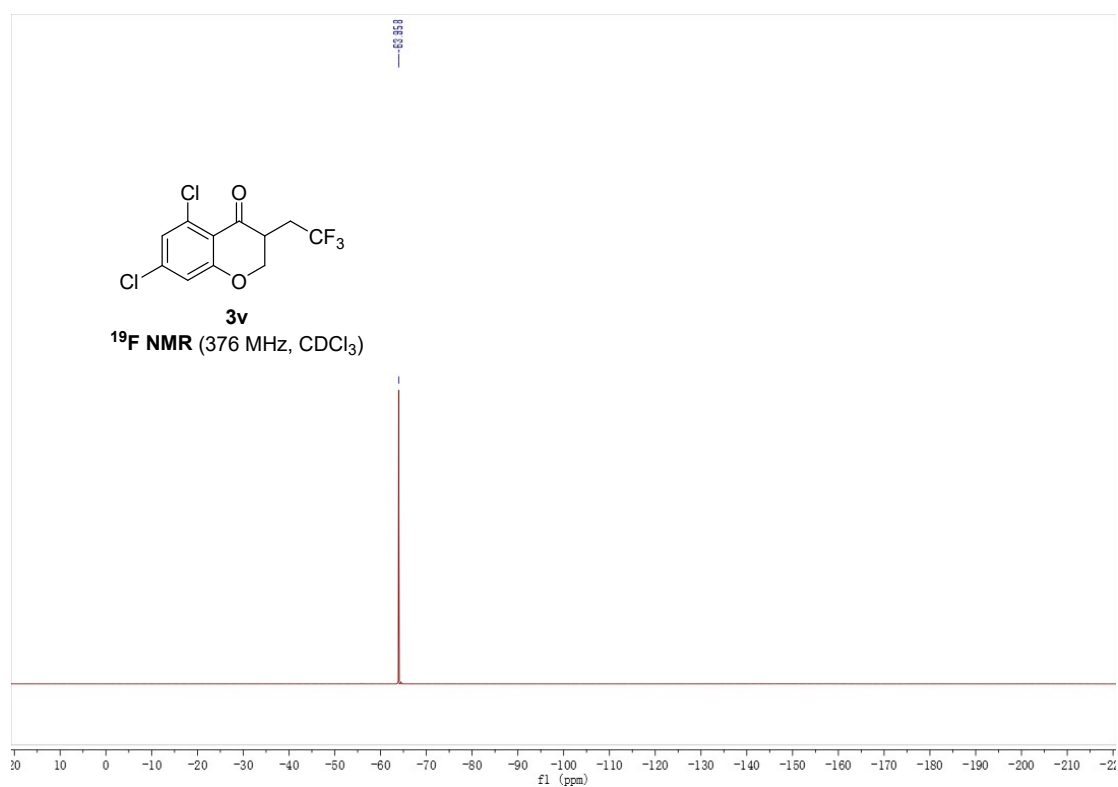
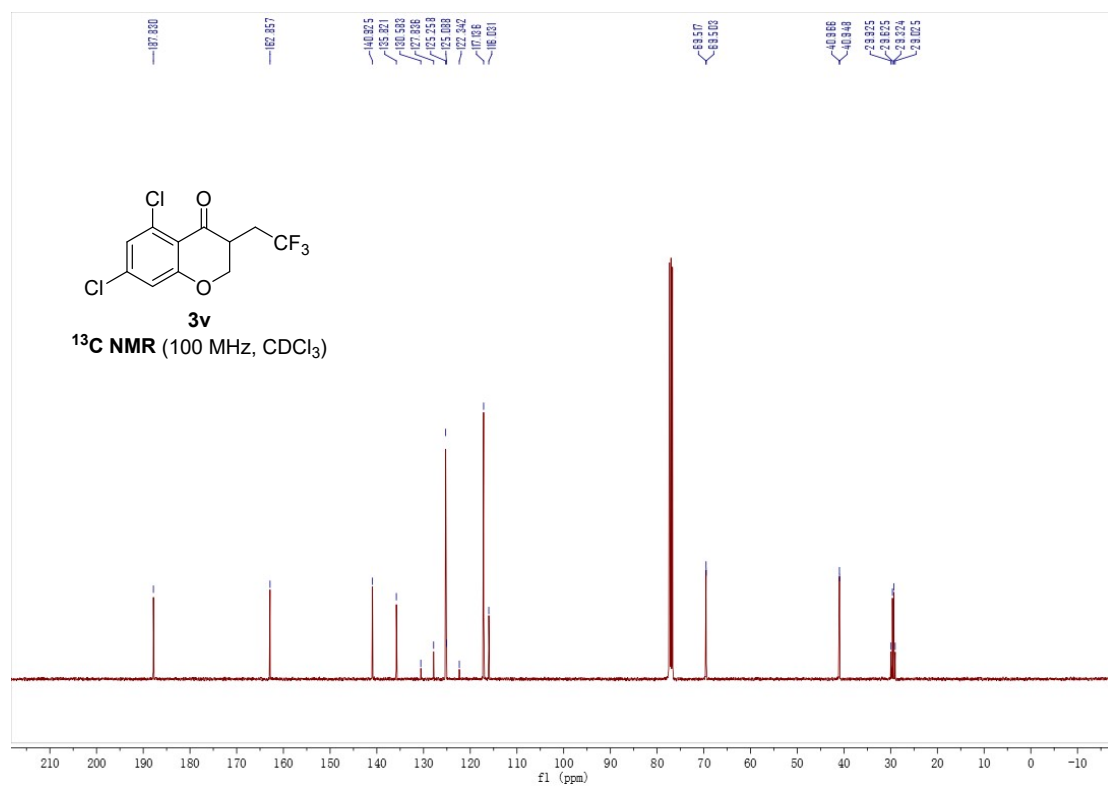
6-bromo-8-fluoro-3-(2,2,2-trifluoroethyl)chroman-4-one (3u)



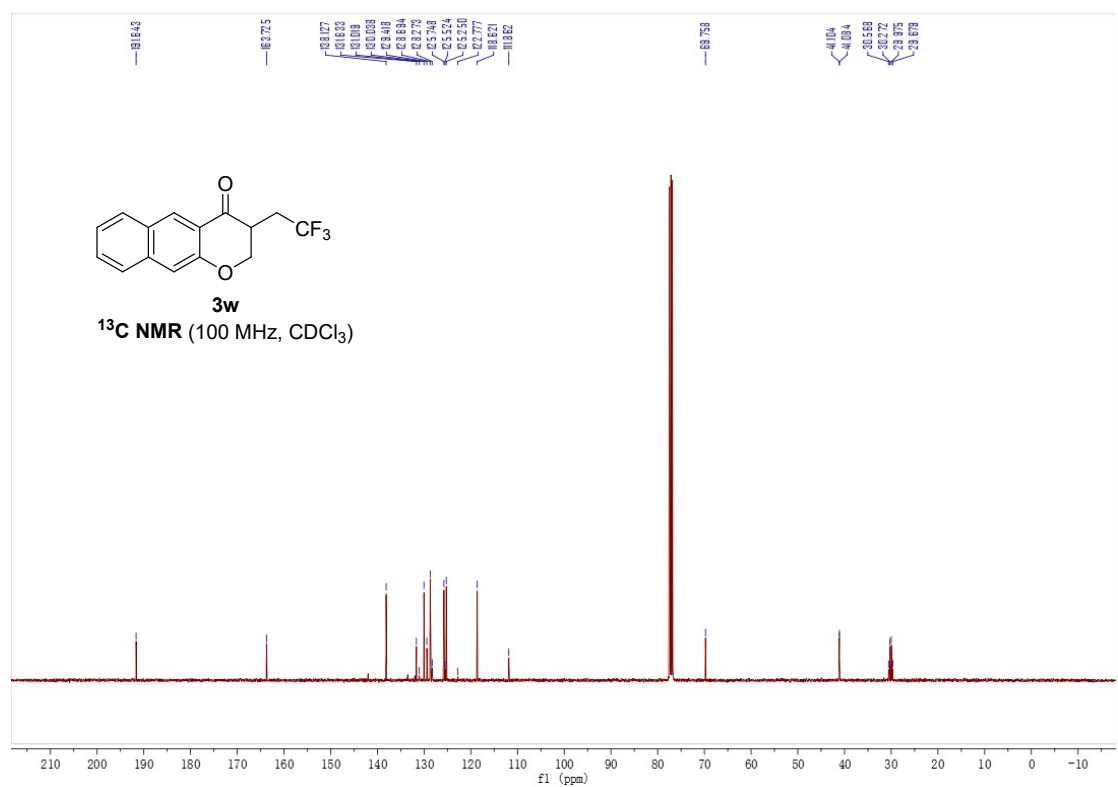
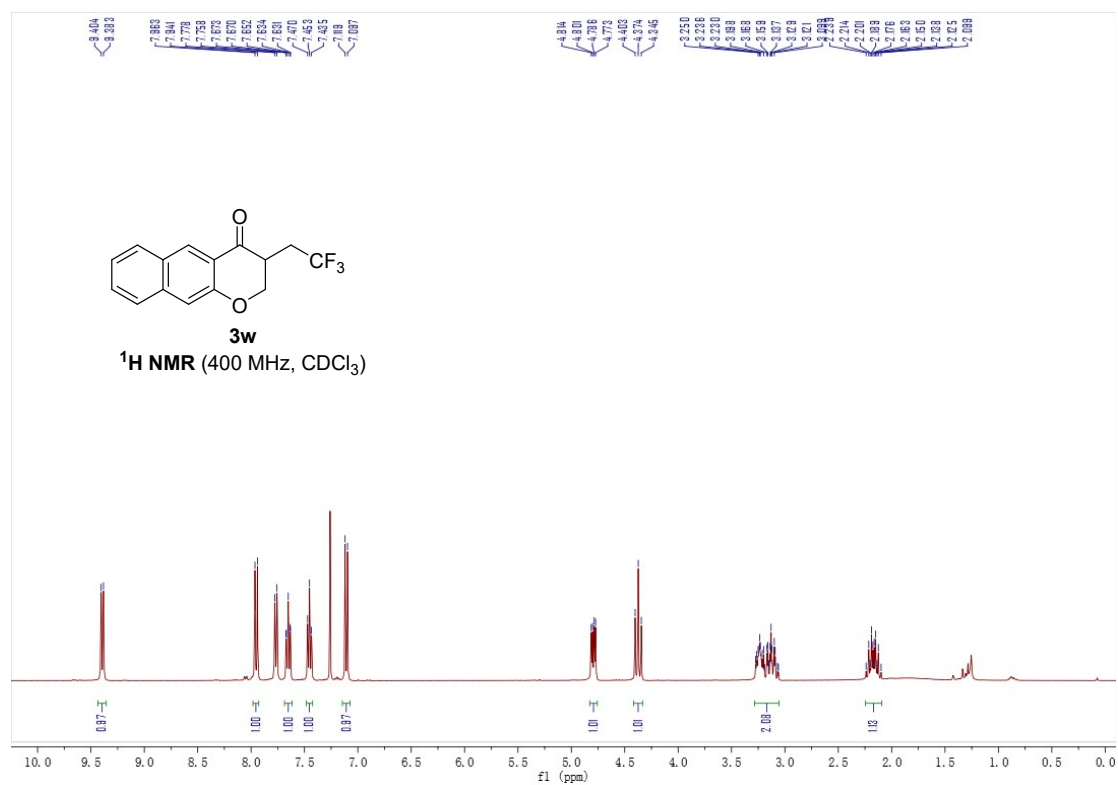


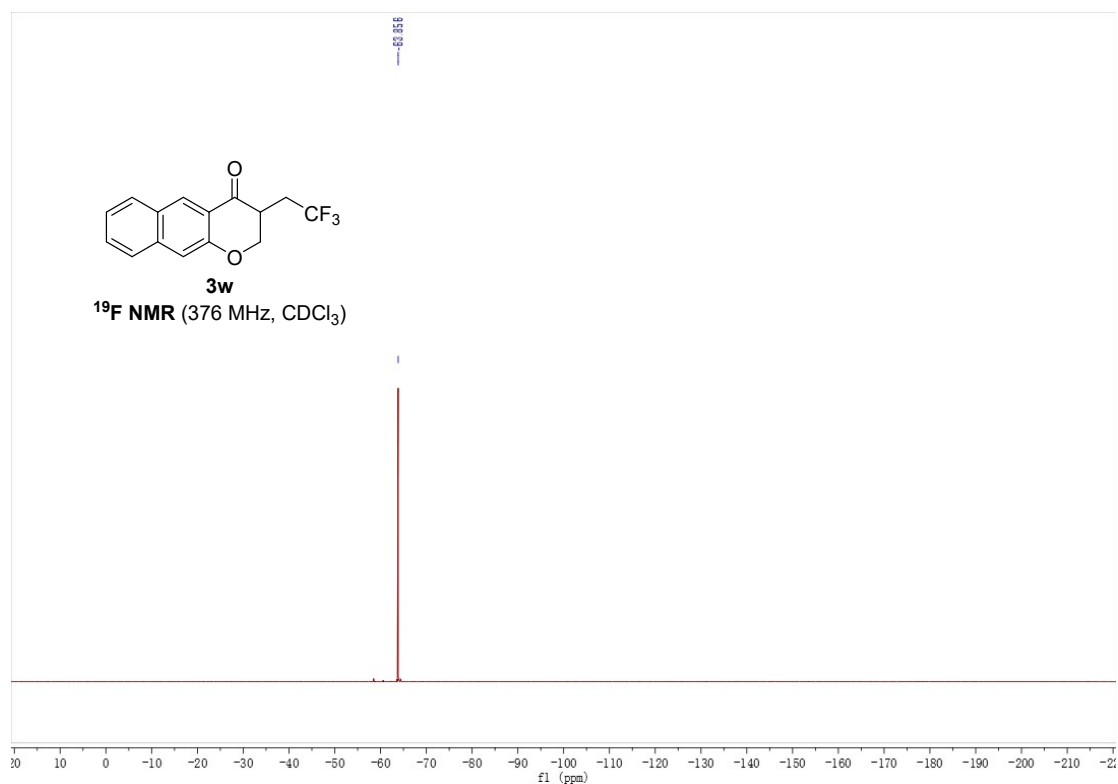
5,7-dichloro-3-(2,2,2-trifluoroethyl)chroman-4-one (3v)



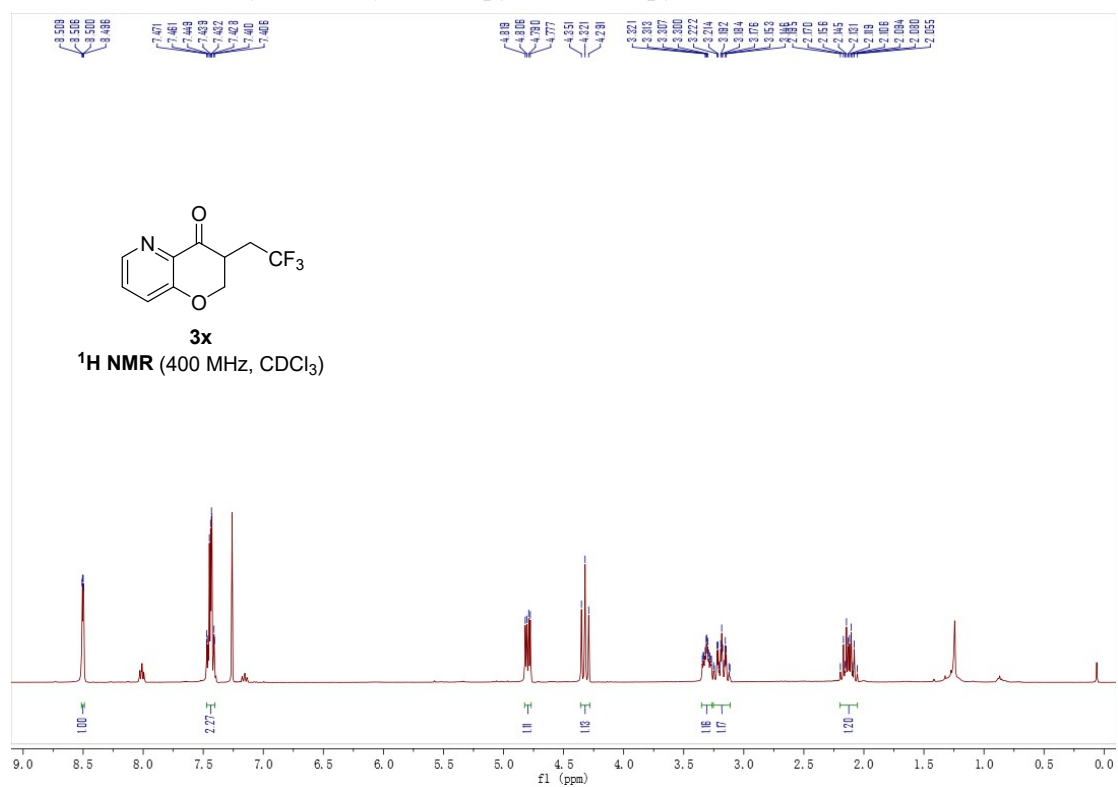


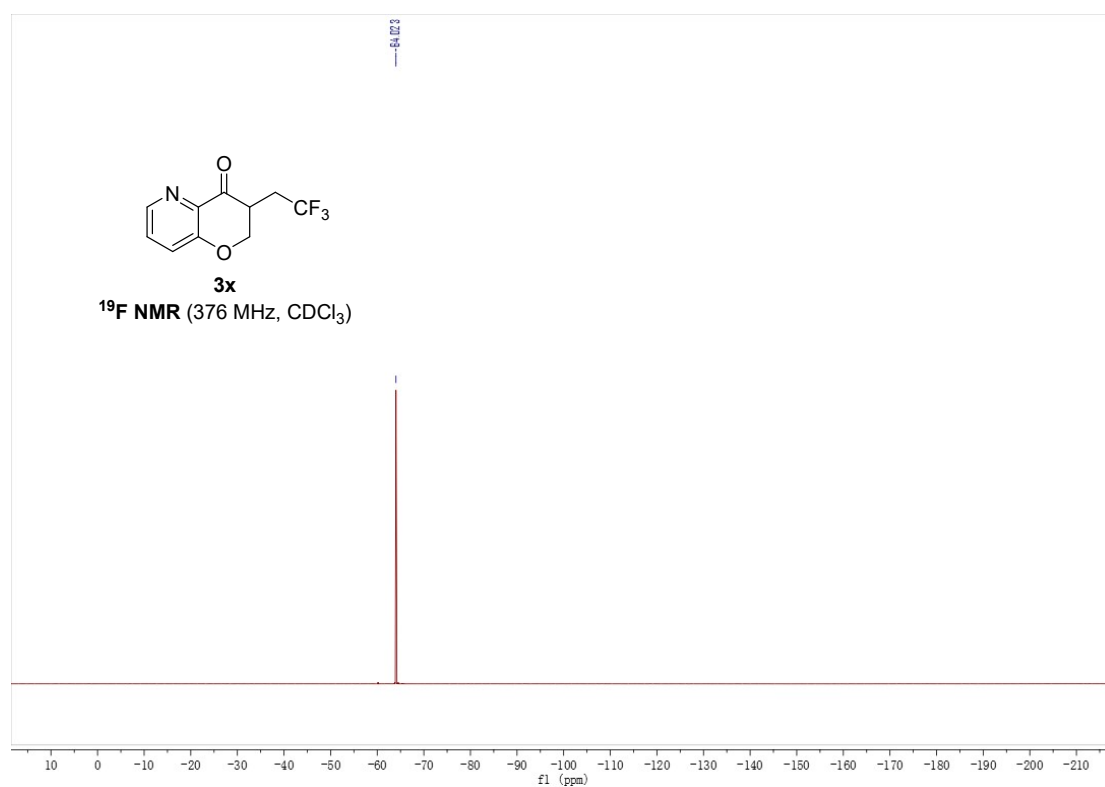
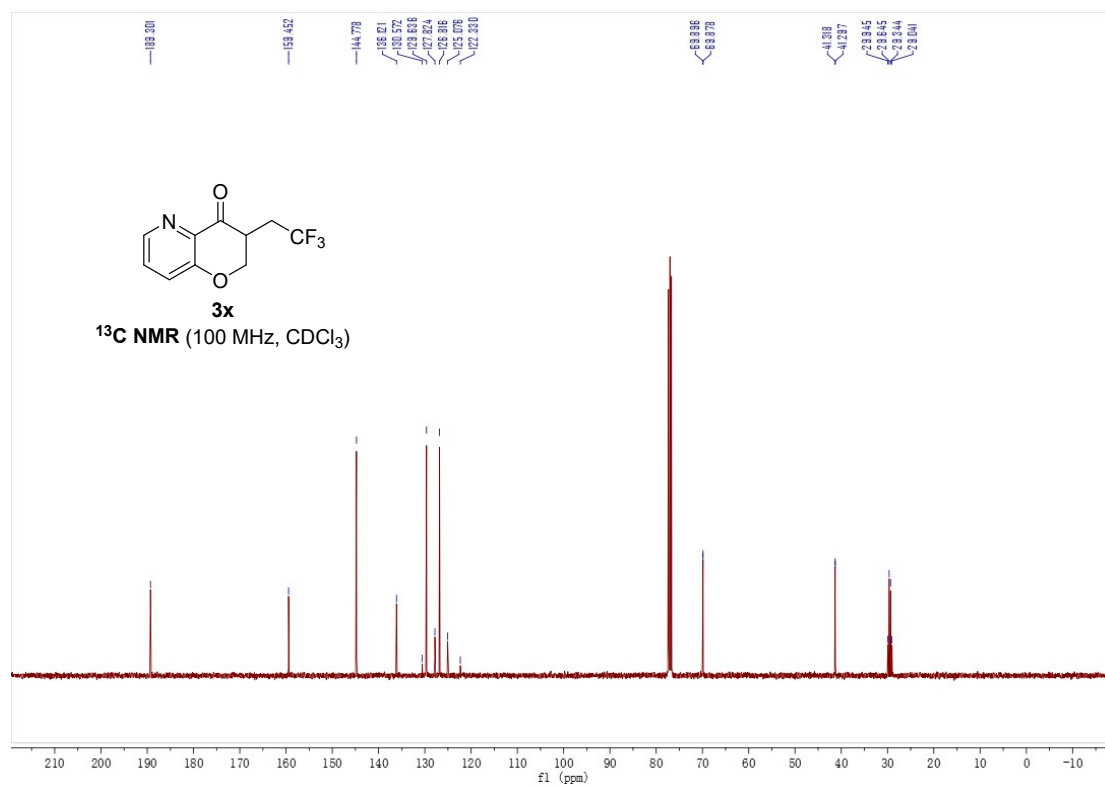
3-(2,2,2-trifluoroethyl)-2,3-dihydro-4H-benzo[g]chromen-4-one (3w)



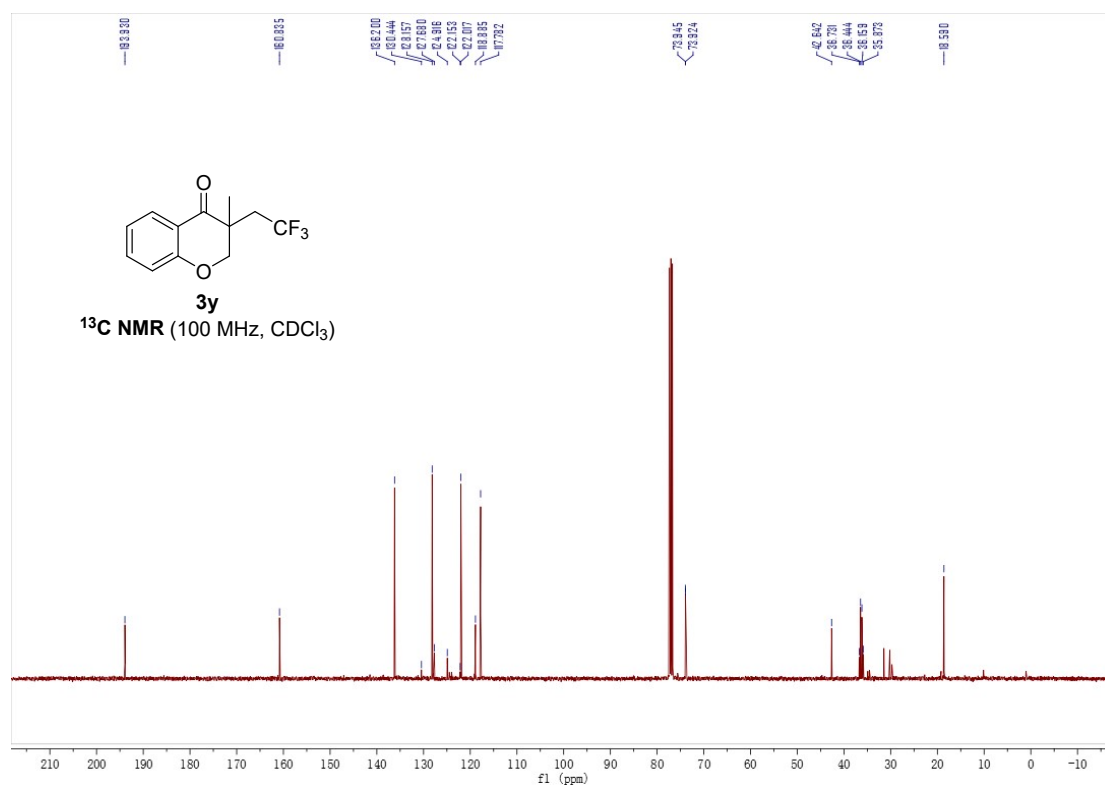
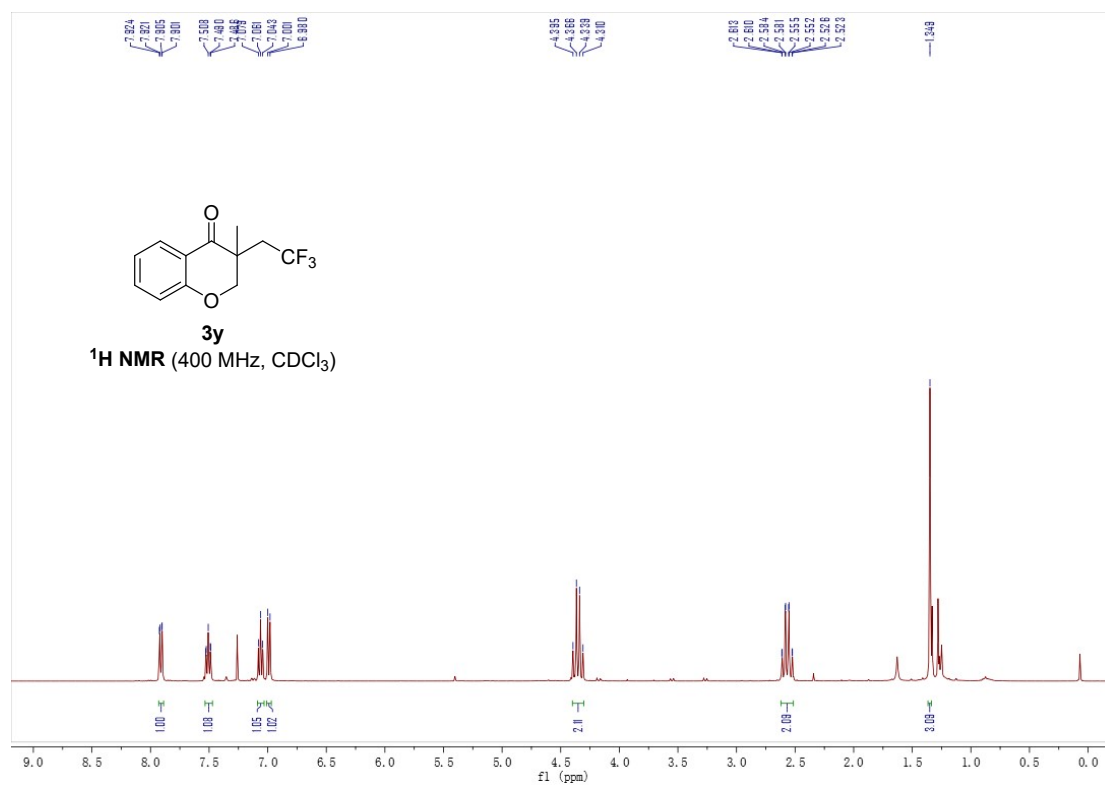


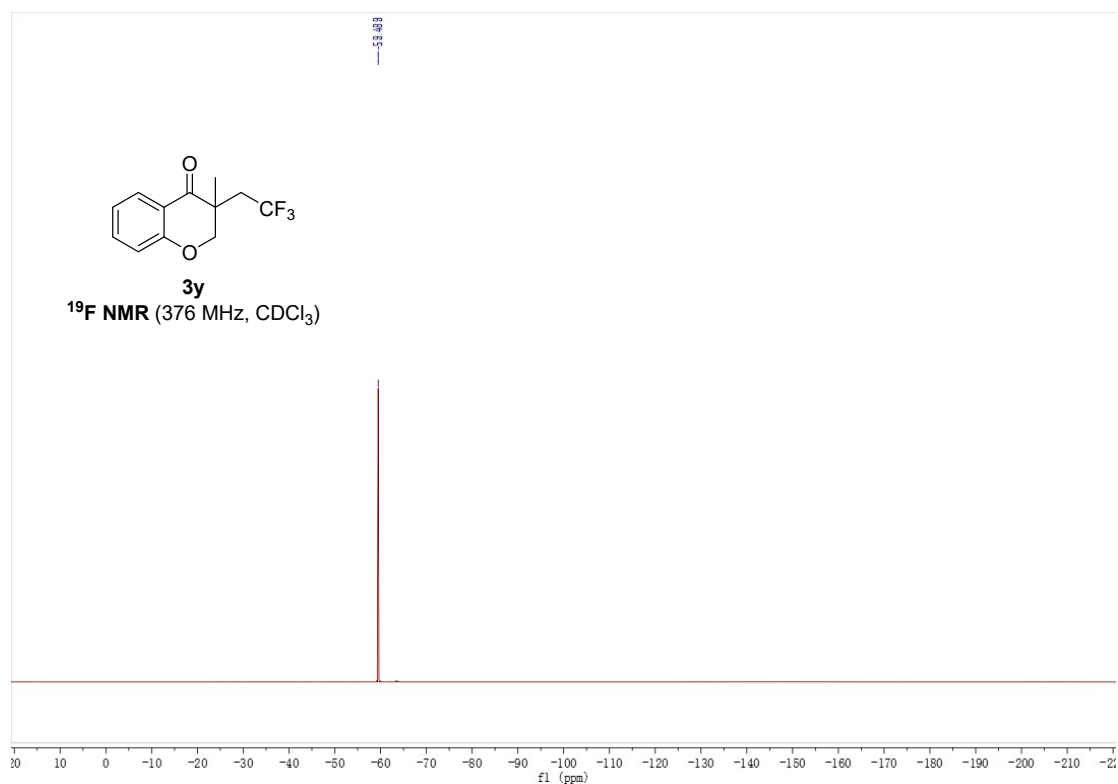
3-(2,2,2-trifluoroethyl)-2,3-dihydro-4H-pyrano[3,2-b]pyridin-4-one (3x)



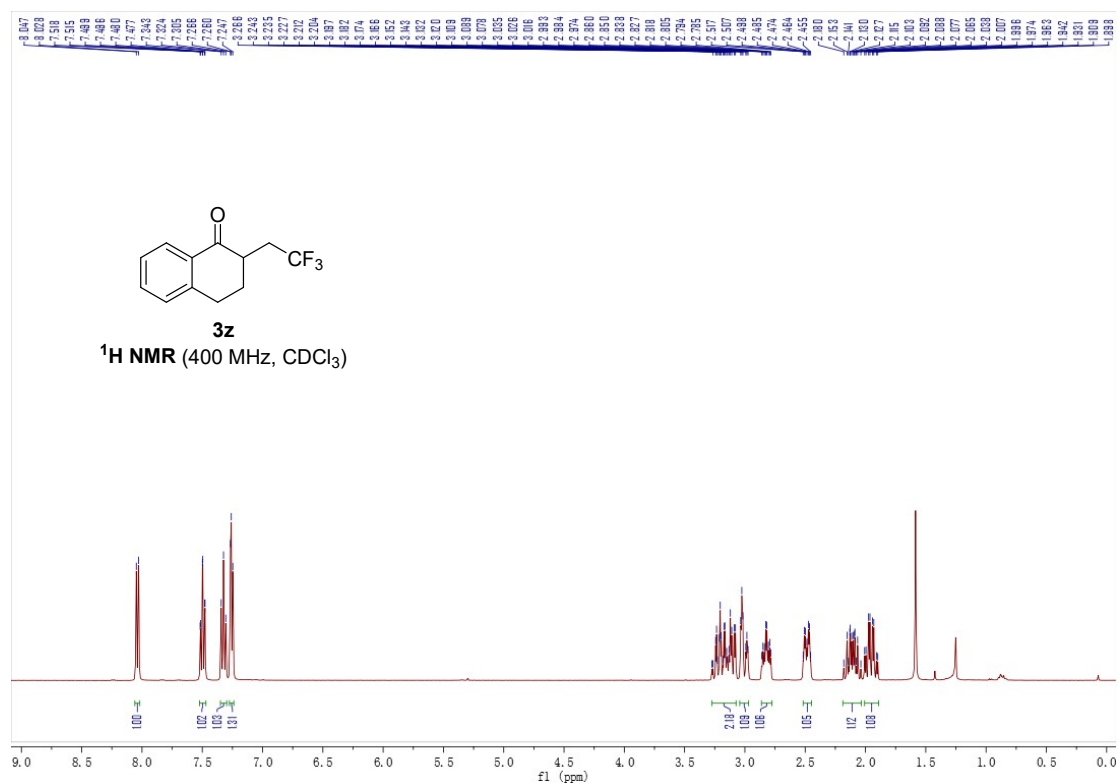


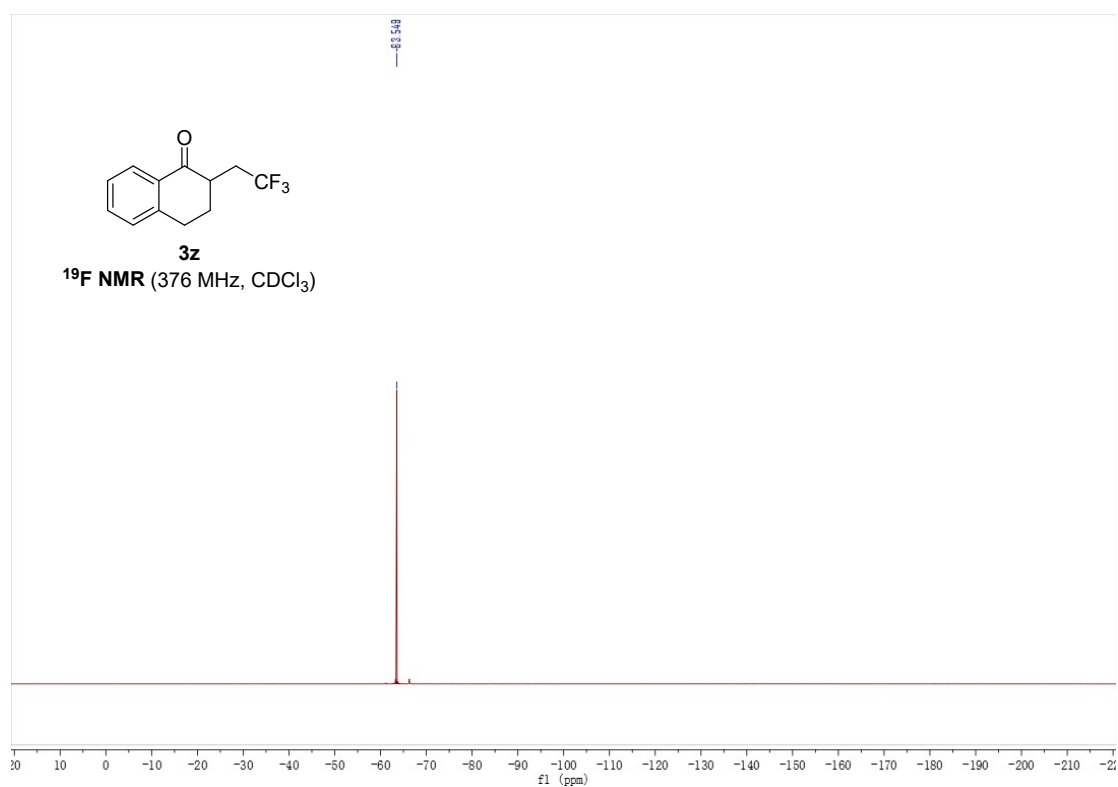
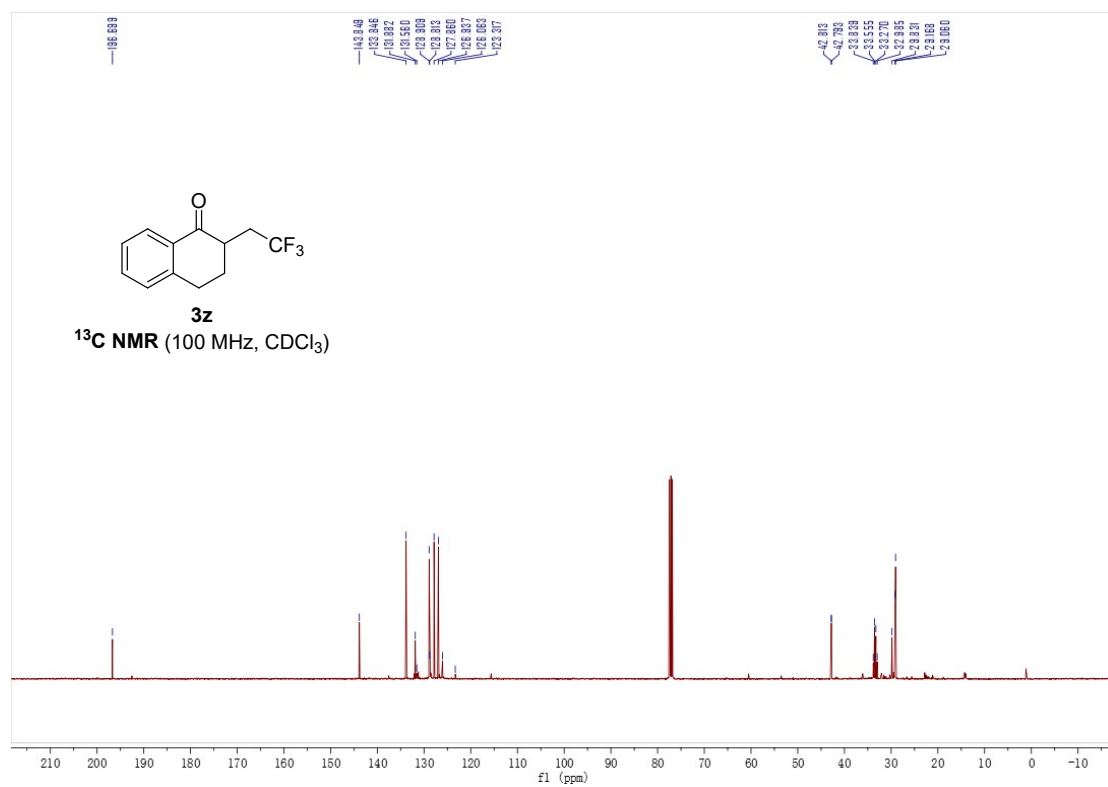
4-methyl-3-(2,2,2-trifluoroethyl)chroman-4-one (3y)



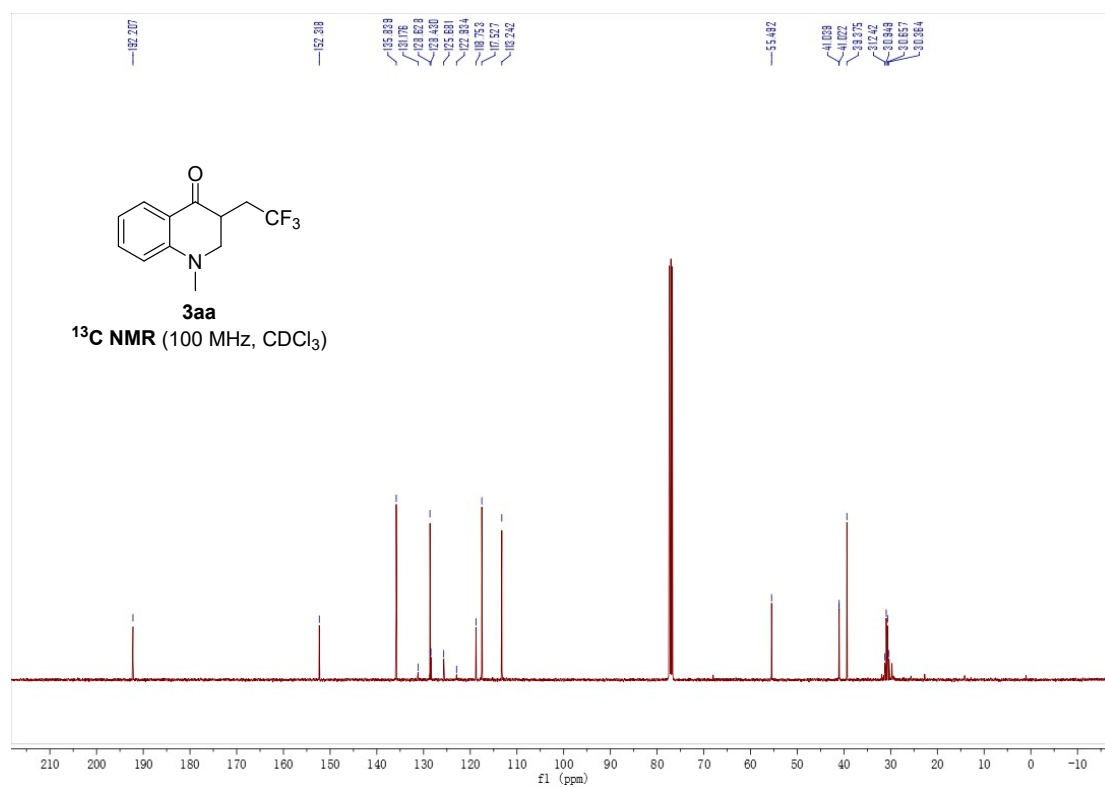
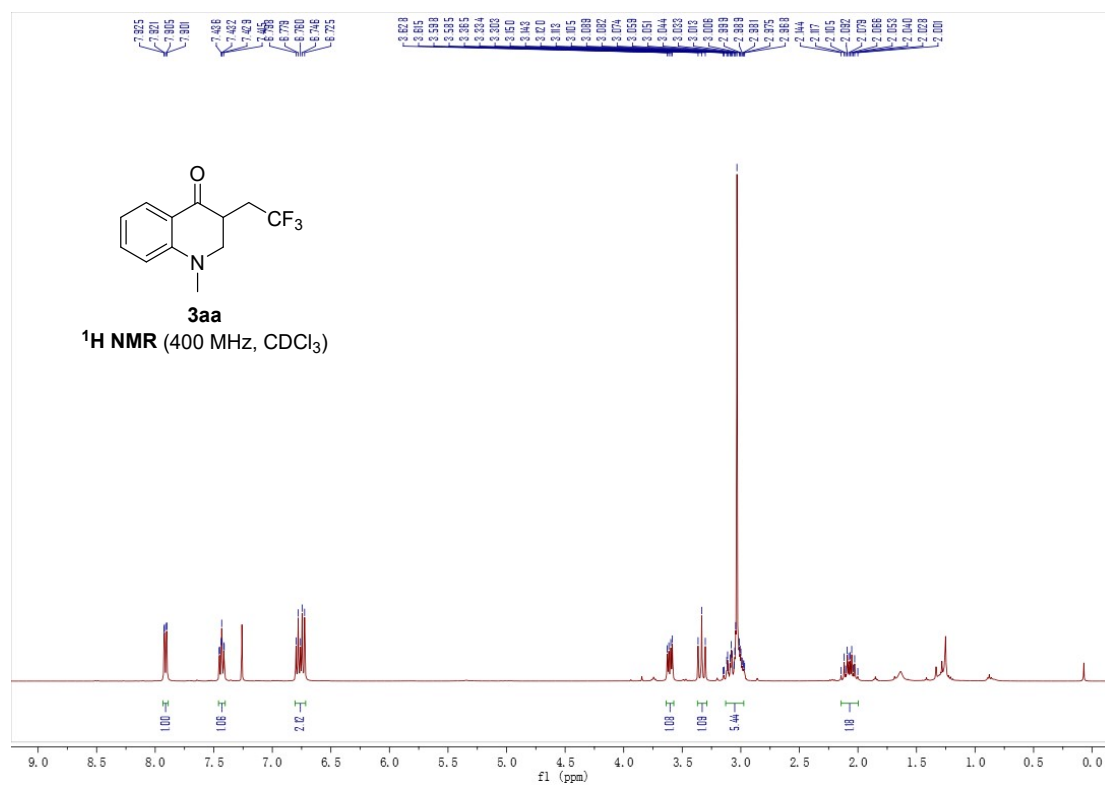


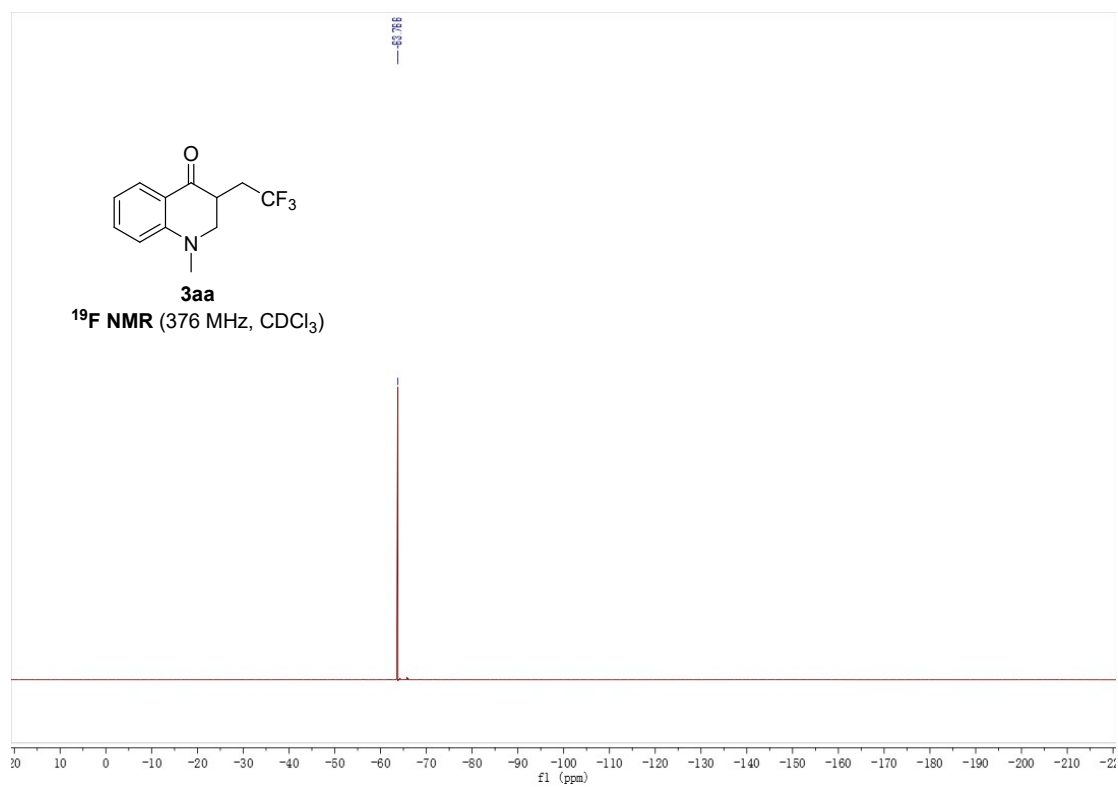
2-(2,2,2-trifluoroethyl)-3,4-dihydronaphthalen-1(2H)-one (3z)



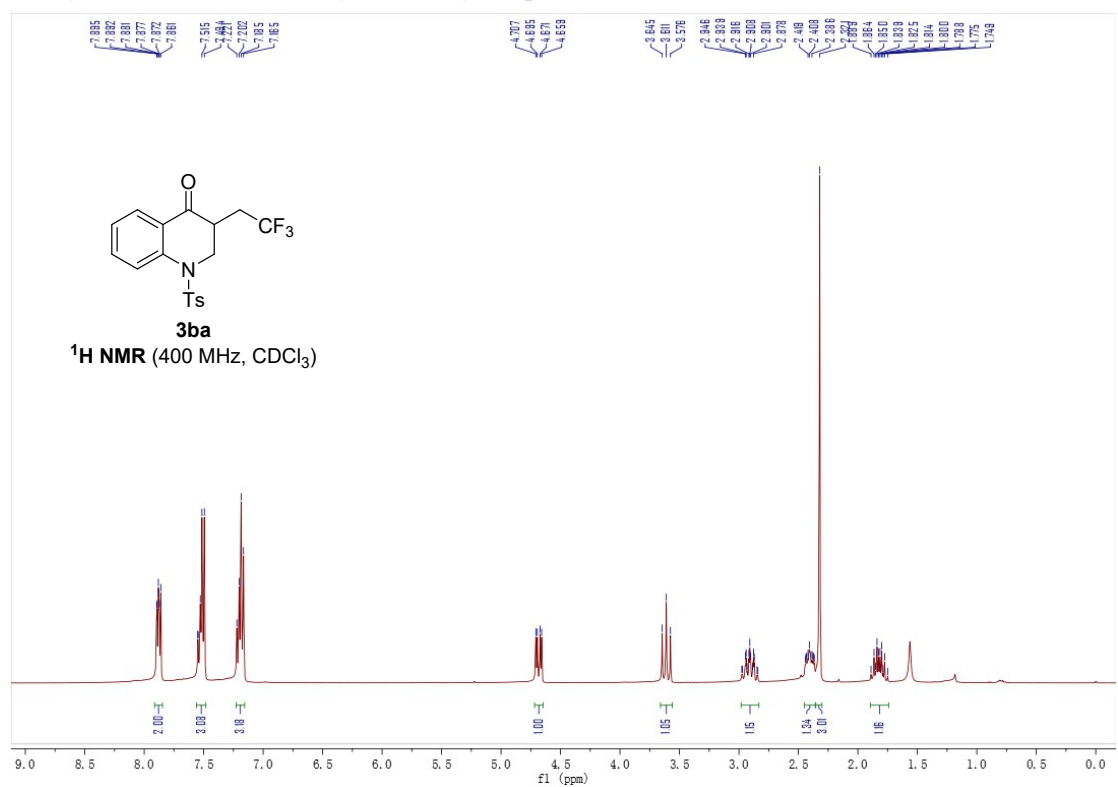


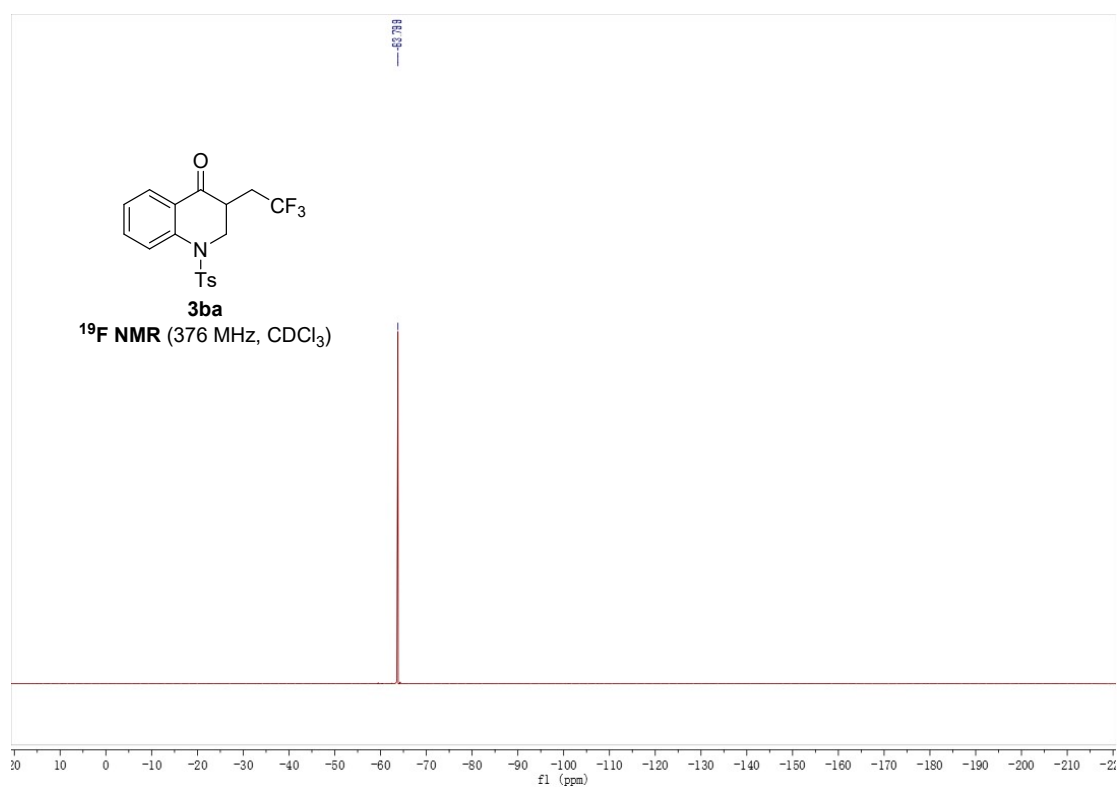
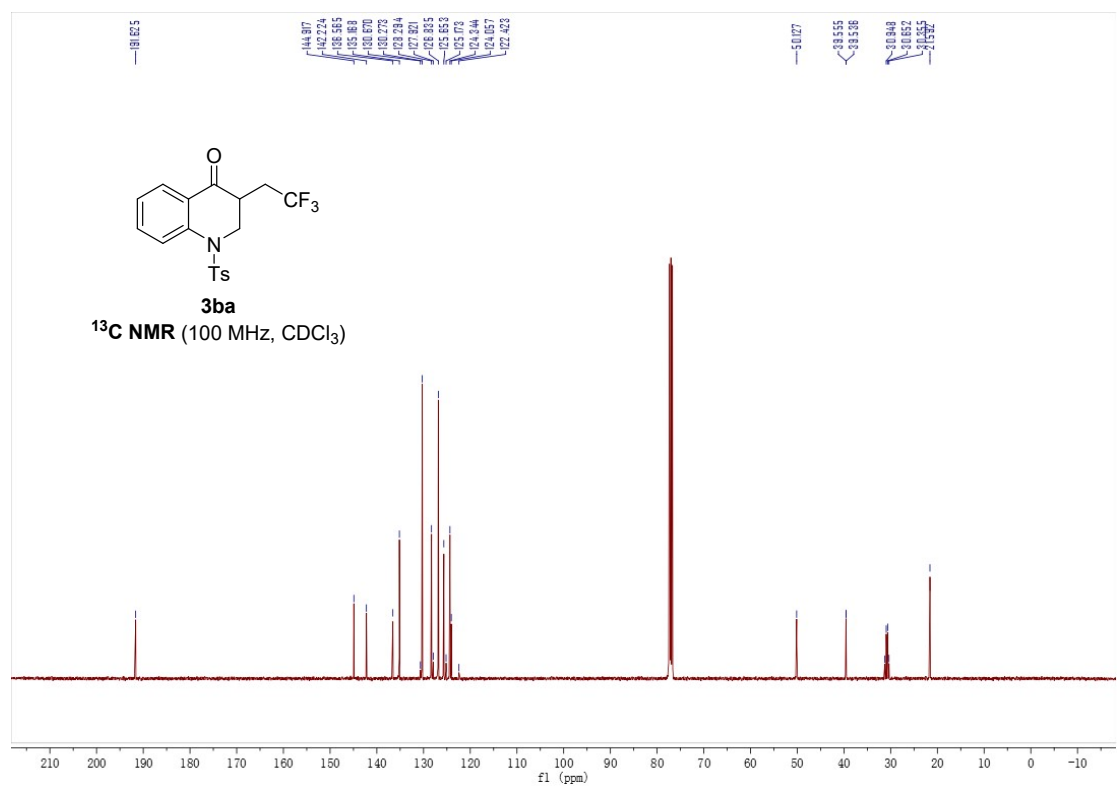
1-methyl-3-(2,2,2-trifluoroethyl)-2,3-dihydroquinolin-4(1H)-one (3aa)



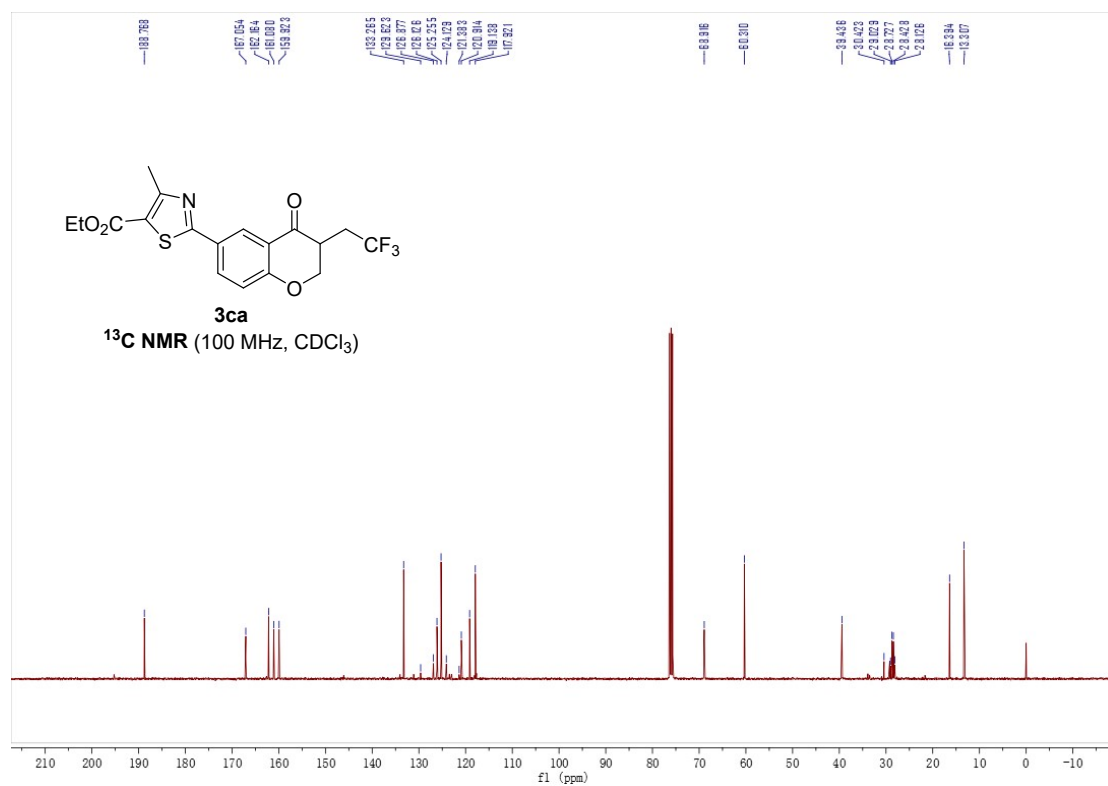
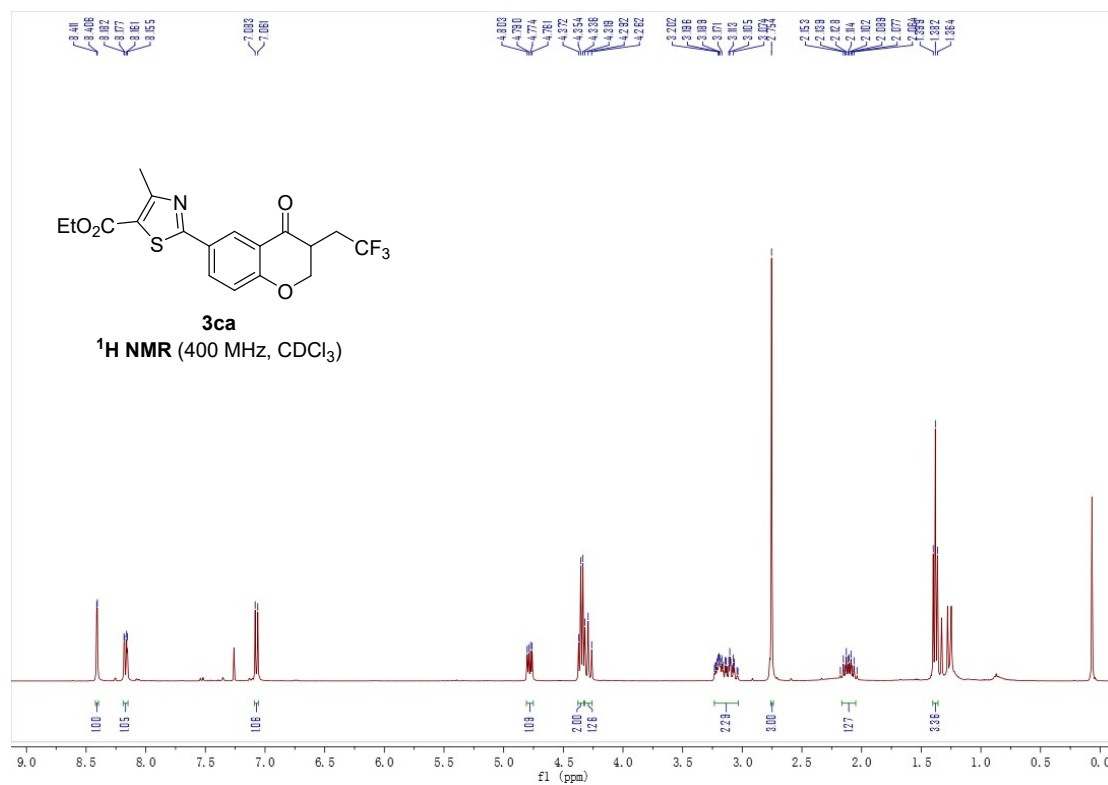


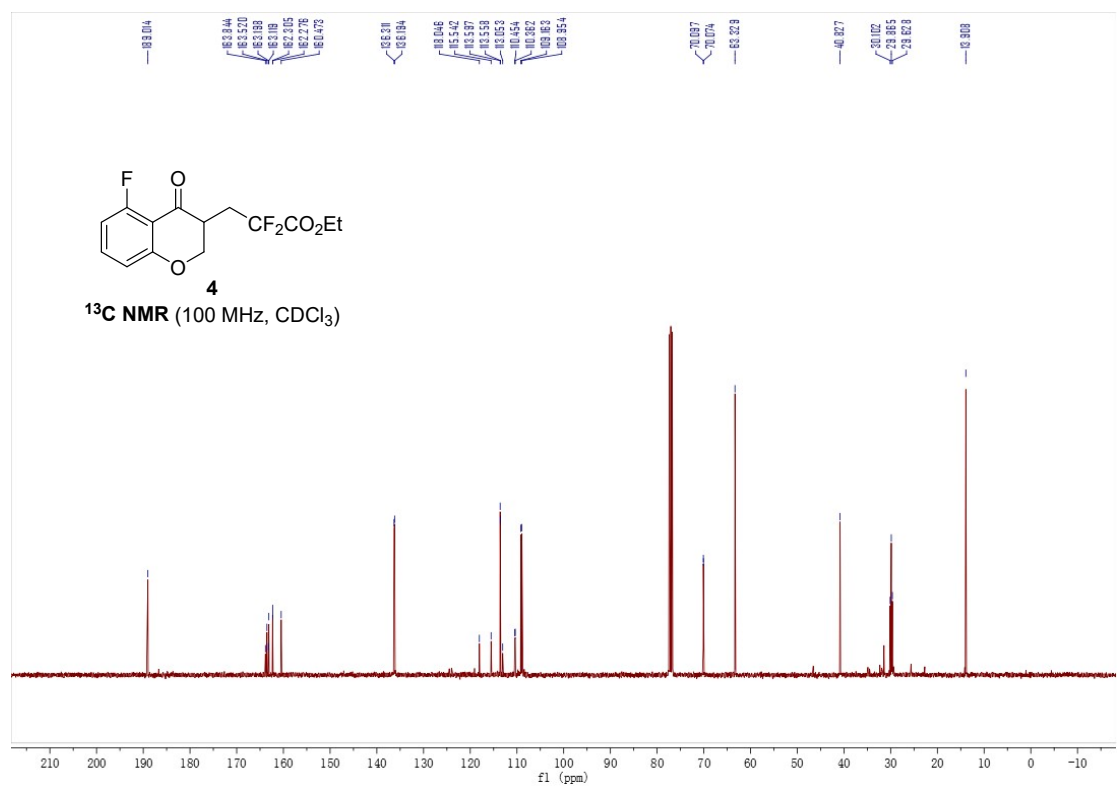
1-tosyl-3-(2,2,2-trifluoroethyl)-2,3-dihydroquinolin-4(1H)-one (3ba)





Ethyl 4-methyl-2-(4-oxo-3-(2,2,2-trifluoroethyl)chroman-6-yl)thiazole-5-carboxylate (3ca)





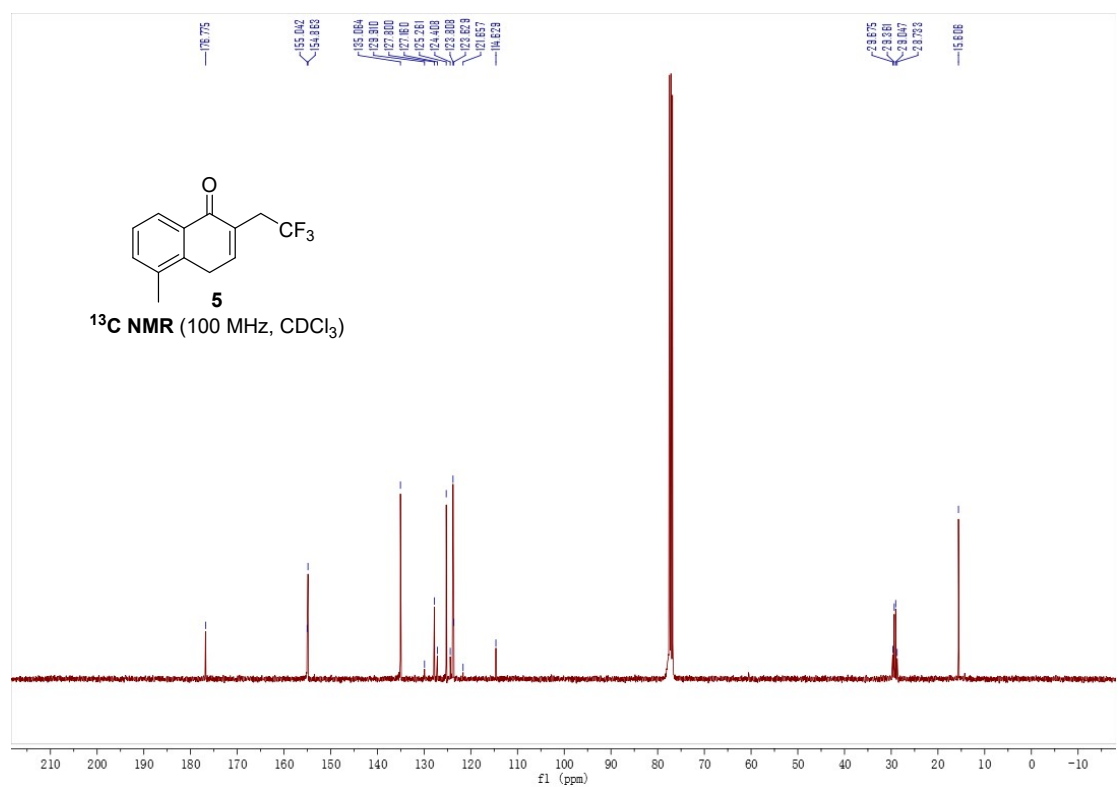
5

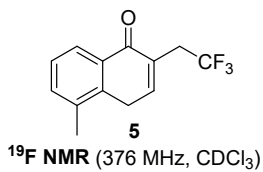
¹H NMR (400 MHz, CDCl₃)

Chemical structure of compound **5** is shown in the top left corner. The structure is a 1,2,3,4-tetrahydronaphthalen-1-one derivative with a 2-(trifluoromethyl)ethyl group at position 2 and a methyl group at position 6.

The ¹H NMR spectrum (400 MHz, CDCl₃) shows the following peaks and integrations:

- Aromatic region (7.2-8.1 ppm): Multiple peaks with integrations of 1.00, 1.00, 1.04, and 1.04.
- Aliphatic region (1.2-1.8 ppm): Multiple peaks with integrations of 2.24 and 3.16.





6
¹H NMR (400 MHz, CDCl₃)

