

Supporting Information to
Cobalt-Promoted Selective Arylation of Benzamides and
Acrylamides with Arylboronic Acids

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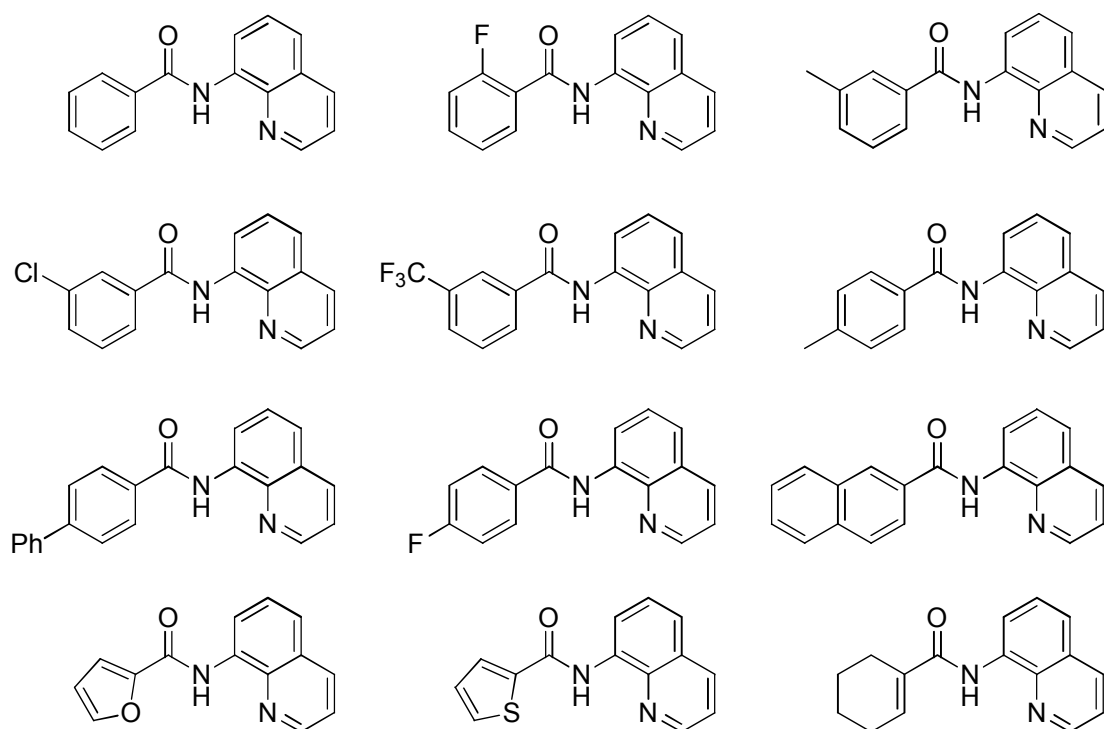
1. General information

^1H NMR, ^{13}C NMR and ^{19}F NMR were recorded in CDCl_3 at room temperature on the Varian INOVA-400 spectrometer (400 MHz, ^1H). The ^1H NMR chemical-shifts scale is based on internal TMS. The peak patterns are indicated as follows: s, singlet; d, doublet; t, triplet; q, quartet; m, multiplet; qui, quintet; sxt, sextet. The coupling constants, J are reported in Hertz (Hz). High-resolution mass spectral (HRMS) analyses were carried out using a TOF MS instrument with an ESI source.

Unless otherwise noted, all reagents were obtained from commercial suppliers and used without further purification. Anhydrous $\text{Co}(\text{acac})_2$ was purchased from Alfa Aesar. All solvents were purified and dried according to standard methods prior to use. Products were purified by flash column chromatography on 200-300 mesh silica gel, SiO_2 .

2. Typical procedure for the preparation of benzamides

All benzamides were synthesized from the corresponding benzoic acids or benzoyl chlorides and 8-aminoquinoline. The deuterated amides were synthesized according to a literature method, spectral properties are consistent with literature values.¹ The following amides were synthesized according to literature procedures.²



3. Co-promoted arylation of arylboronic acids

3.1 Optimization of reaction conditions

Scheme S1 ineffective directing groups

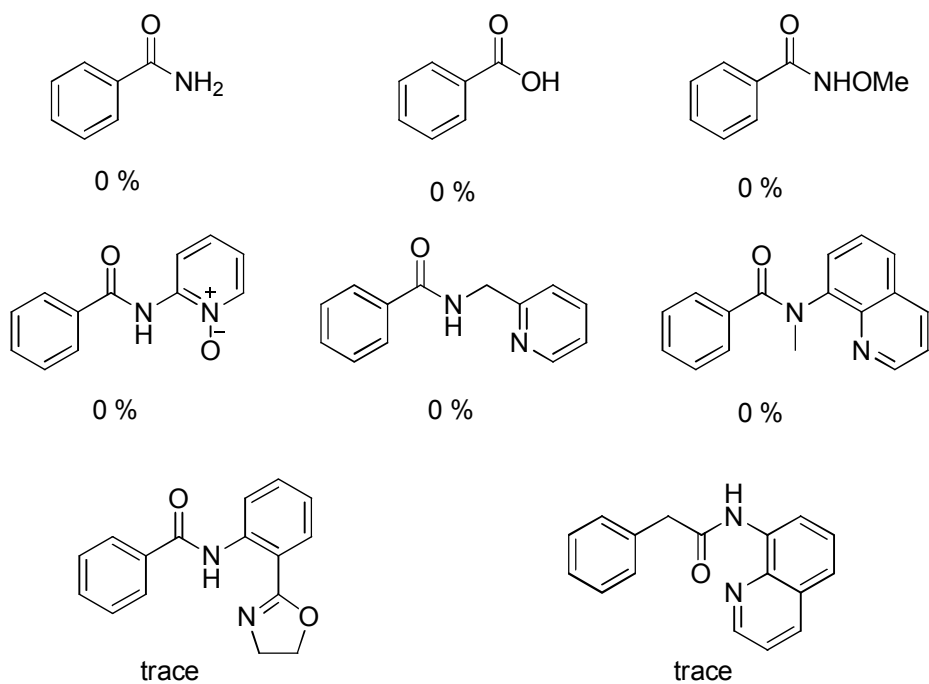


Table S1. screening of Co salts ^a

Entry	Co salts	yield ^b (%)
1	Co(OAc) ₂	trace
2	Co(CO) ₃	trace
3	CoCl ₂	ND
4	CoBr ₂	ND
5	CoSO ₄	ND
6	Co(oxalate) ₂	trace
7	Co(NO ₃) ₂ ·6H ₂ O	ND
8	Co(acac) ₃	21

^a Reaction conditions: amide **1a** (0.2 mmol), **2a** (0.4 mmol), Co salt (0.2 mmol), Ag₂CO₃ (0.4 mmol), Na₂CO₃ (0.2 mmol), DMSO (1 mL), 120 °C, under air for 2h; ^b isolated yield.

Table S2. screening of oxidant ^a

Entry	oxidant	yield ^b (%)
1	Mn(OAc) ₂	17
2	NMO	ND
3	O ₂ (1 atm)	ND
4	Ag ₂ O	52
5	AgOAc	21
6	AgCF ₃ CO ₂	14

^a Reaction conditions: amide **1a** (0.2 mmol), **2a** (0.4 mmol), Co(acac)₂ (0.2 mmol), oxidant (0.4 mmol), Na₂CO₃ (0.2 mmol), DMSO (1 mL), 120 °C, under air for 2h; ^b isolated yield.

Table S3. screening of base ^a

Entry	base	yield ^b (%)
1	K ₂ CO ₃	72
2	NaOPiv	53
3	NaHCO ₃	82
4	Cs ₂ CO ₃	54
5	<i>t</i> -BuONa	43

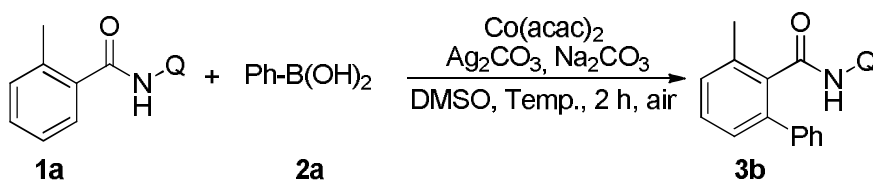
^a Reaction conditions: amide **1a** (0.2 mmol), **2a** (0.4 mmol), Co(acac)₂ (0.2 mmol), Ag₂CO₃ (0.4 mmol), base (0.2 mmol), DMSO (1 mL), 120 °C, under air for 2h; ^b isolated yield.

Table S4. screening of solvent ^a

Entry	solvent	yield ^b (%)
1	DMF	79
2	NMP	64
3	<i>t</i> -BuOH	ND
4	MeCN	trace
5	toluene	trace
6	dioxane	ND
7	DCE	ND

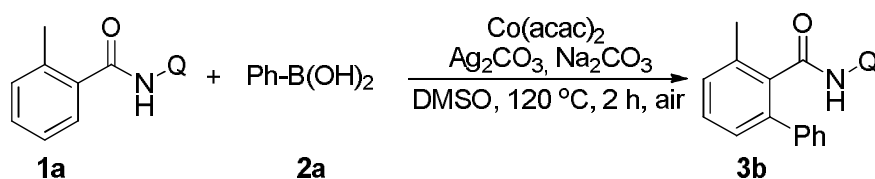
^a Reaction conditions: amide **1a** (0.2 mmol), **2a** (0.4 mmol), Co(acac)₂ (0.2 mmol), Ag₂CO₃ (0.4 mmol), Na₂CO₃ (0.2 mmol), solvent (1 mL), 120 °C, under air for 2h; ^b isolated yield.

Table S5. screening of temperature ^a

		
Entry	Temp. (°C)	yield ^b (%)
1	R.T.	trace
2	60	<10
3	80	29
4	100	56
5	120	91
6	140	81

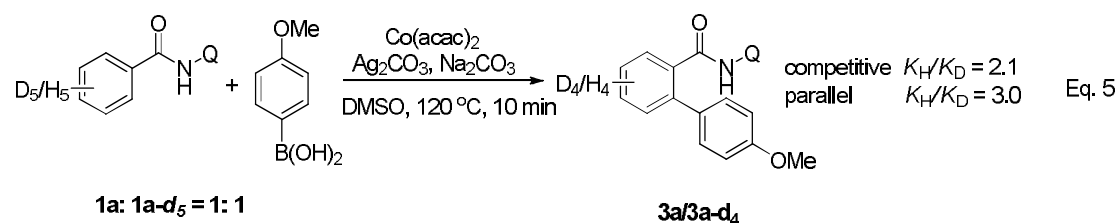
^a Reaction conditions: amide **1a** (0.2 mmol), **2a** (0.4 mmol), Co(acac)₂ (0.2 mmol), Ag₂CO₃ (0.4 mmol), Na₂CO₃ (0.2 mmol), DMSO (1 mL), under air for 2h; ^b isolated yield.

3.2 General procedure for Co-Promoted *ortho*-Arylation of Benzamides with Aryboronic Acids:



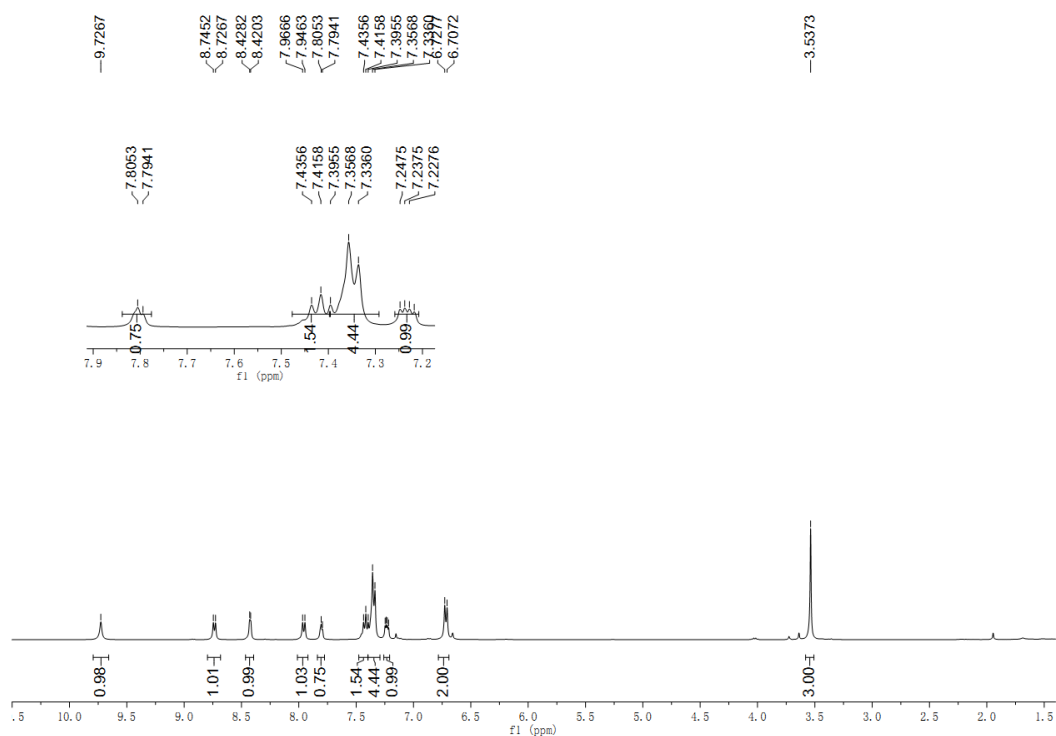
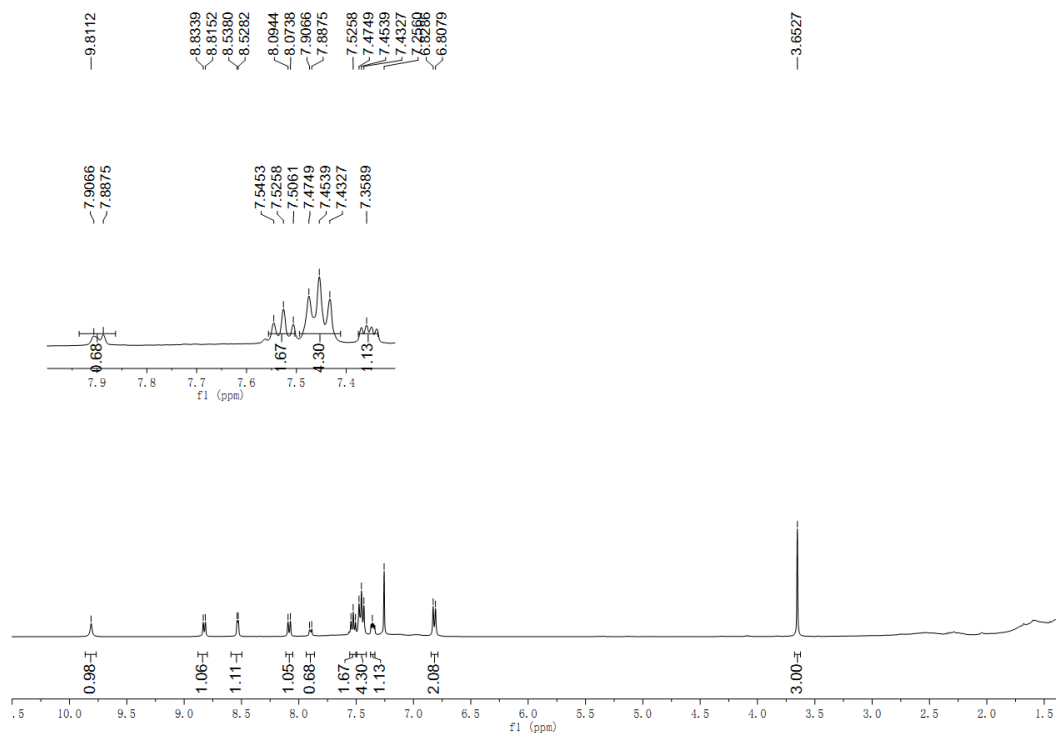
Benzamide **1a** (0.2 mmol), anhydrous Co(acac)₂ (49 mg, 0.2 mmol), Ag₂CO₃ (110 mg, 0.4 mmol), Na₂CO₃ (21 mg, 0.2 mmol), arylboronic acid **2a** (0.4 mmol) and anhydrous DMSO (1 mL) were added to a 25-mL Schlenk flask equipped with a high-vacuum PTFE valve-to-glass seal. Then the flask was sealed under air and stirred at 120 °C for 2 h. After the completion of the reaction, the solvent was evaporated under reduced pressure. The mixture was extracted with ethyl acetate, and the combined organic layer was dried over sodium sulfate. Concentration in vacuo followed by silica gel column purification with petroleum ether/ethyl acetate eluent (8/1 to 6/1) gave the desired product **3a** as a colorless oil (62 mg, 91% yield).

3.3 Deuterium-labeling experiments:

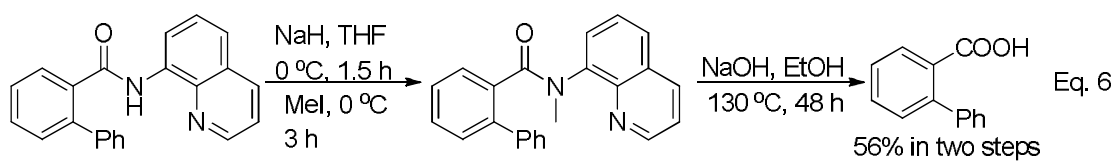


Intermolecular competition KIE Following general procedure: **1a** (50 mg, 0.2 mmol), **1a'**-**d₅** (51 mg, 0.2 mmol), Co(acac)₂ (98 mg, 0.4 mmol), Ag₂CO₃ (220 mg, 0.8 mmol), Na₂CO₃ (42 mg, 0.4 mmol), **2h** (122 mg, 0.8 mmol) and anhydrous DMSO (2 mL) were added to a 25-mL Schlenk flask equipped with a high-vacuum PTFE valve-to-glass seal. Then the flask was sealed under air and stirred at 120 °C for 10 min. The product was separated by column chromatography to give the desired product less than 13 % yield. The KIE value was calculated as $k_{\text{H}}/k_{\text{D}} = 2.1$.

Intermolecular parallel KIE Following general procedure: **1a** (50 mg, 0.2 mmol) or **1a'**-**d₅** (51 mg, 0.2 mmol), Co(acac)₂ (49 mg, 0.2 mmol), Ag₂CO₃ (110 mg, 0.4 mmol), Na₂CO₃ (21 mg, 0.2 mmol), **2h** (61 mg, 0.4 mmol) and anhydrous DMSO (1 mL) were added to a 25-mL Schlenk flask equipped with a high-vacuum PTFE valve-to-glass seal. Then the flask was sealed under air and stirred at 120 °C for 10 min. The product was separated by column chromatography to give the desired product less than 16 % yield. The KIE value was calculated as $k_{\text{H}}/k_{\text{D}} = 3.0$.



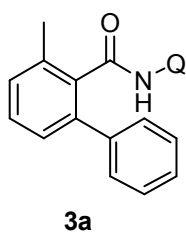
3.4 Removal of the 8-aminoquinoline Auxiliary:



N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide **4a** (324 mg, 1 mmol) was dissolved in anhydrous THF (10 mL) and the resulting solution was cooled to 0°C. To this solution, NaH (80 mg, 2 mmol) was added in portions over 10 min. The resulting solution was allowed to stir for 1.5 h. MeI (710 mg, 5 mmol) was added dropwise over 5 min, and reaction mixture was stirred for additional 3 hours at 0 °C and stirred overnight at rt. After the reaction was quenched by addition water, the mixture was extracted with Et₂O, and the organic layer was dried by anhydrous Na₂SO₄. After remove the solvent, the residue was purified by column chromatography to give intermediate **7** as a yellow solid (271 mg, 80%). Intermediate **7** (169 mg, 0.5 mmol) and NaOH (307 mg, 7.5 mmol) were dissolved in EtOH (5 ml). The resulting mixture was stirred at 130 °C for 48 hours. After that, reaction mixture was diluted with EtOAc (100 mL) and 1N HCl (30mL) was added. Organic layer was washed with 1N HCl (5 x 20 mL), dried over anhydrous Na₂SO₄, filtered and the solvent was evaporated under vacuum affording pure product as a faint yellow solid (56%). Its spectral properties are consistent with literature values.¹⁶

4 Characterization data of products

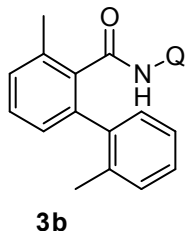
3-Methyl-*N*-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**3a**)



91% yield (62 mg); colorless oil; ¹H NMR (400 MHz, CDCl₃) δ 9.65 (s, 1H), 8.77 (d, *J* = 7.5 Hz, 1H), 8.55 (d, *J* = 4.0 Hz, 1H), 7.98 (d, *J* = 8.2 Hz, 1H), 7.53–7.51 (m, 2H), 7.49–7.45 (m, 1H), 7.41–7.36 (m, 2H), 7.31–7.26 (m, 3H), 7.19 (t, *J* = 7.7 Hz, 2H), 7.06 (t, *J* = 7.5 Hz, 1H), 2.52 (s, 3H); ¹³C NMR (100 MHz, CDCl₃) δ 168.3, 147.9, 140.3, 139.6, 138.3, 136.7, 136.0, 135.7, 134.2, 129.4, 129.2, 128.6, 128.1, 127.7, 127.6, 127.2, 127.1, 121.7, 121.4, 116.4, 19.7; HRMS (ESI) calcd for C₂₃H₁₉N₂O (M + H)⁺ 339.1497,

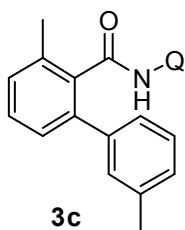
found 339.1488.

2'-Methyl-3-methyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**3b**)



77% yield (54 mg); colorless oil; ^1H NMR (400 MHz, CDCl_3) δ 9.64 (s, 1H), 8.67 (d, $J = 4.6$ Hz, 2H), 8.03 (d, $J = 8.0$ Hz, 1H), 7.41–7.33 (m, 4H), 7.30–7.27 (m, 2H), 7.14 (d, $J = 7.5$ Hz, 1H), 7.05–7.03 (m, 1H), 6.98–6.96 (m, 2H), 2.52 (s, 3H), 2.29 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 167.7, 147.8, 139.6, 139.2, 138.2, 137.4, 136.0, 135.8, 135.6, 134.2, 129.8, 129.6, 129.3, 128.7, 127.7, 127.6, 127.4, 127.2, 125.1, 121.4, 116.1, 20.3, 19.8; HRMS (ESI) calcd for $\text{C}_{24}\text{H}_{21}\text{N}_2\text{O}$ ($\text{M} + \text{H}$) $^+$ 353.1854, found 353.1857.

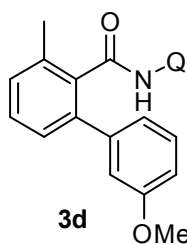
3'-Methyl-3-methyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**3c**)



71% yield (50 mg); colorless oil; ^1H NMR (400 MHz, CDCl_3) δ 9.63 (s, 1H), 8.76 (d, $J = 7.5$ Hz, 1H), 8.59 (d, $J = 4.0$ Hz, 1H), 8.03 (d, $J = 8.2$ Hz, 1H), 7.50–7.46 (m, 1H), 7.43–7.41 (m, 1H), 7.38–7.36 (m, 1H), 7.34–7.26 (m, 5H), 7.06 (t, $J = 7.6$ Hz, 1H), 6.86 (d, $J = 7.5$ Hz, 1H), 2.53 (s, 3H), 2.16 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 168.3, 147.9, 140.3, 139.8, 138.3, 137.6, 136.8, 136.0, 135.7, 134.4, 129.4, 129.3, 129.1, 128.0, 127.9, 127.7, 127.5, 127.2, 125.6, 121.6, 121.4, 116.3, 21.2, 19.8; HRMS (ESI) calcd for $\text{C}_{24}\text{H}_{21}\text{N}_2\text{O}$ ($\text{M} + \text{H}$) $^+$ 353.1854, found 353.1848.

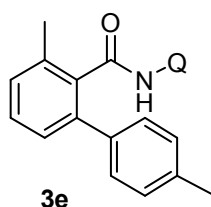
3'-Methoxy-3-methyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**3d**)

73% yield (54 mg); White solid, mp: 137–139 $^\circ\text{C}$ (from ethyl acetate/petroleum ether = 10:1); ^1H NMR (400 MHz, CDCl_3) δ 9.66 (s, 1H), 8.79 (dd, $J = 7.5, 1.3$ Hz, 1H), 8.59 (dd, $J = 4.2, 1.6$ Hz, 1H), 8.04 (dd, $J = 8.3,$



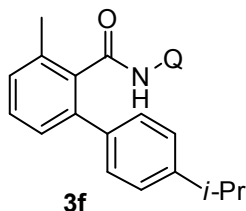
1.5 Hz, 1H), 7.50 (t, $J = 7.9$ Hz, 1H), 7.44–7.37 (m, 2H), 7.34–7.30 (m, 3H), 7.10–7.09 (m, 3H), 6.63–6.60 (m, 1H), 3.63 (s, 3H), 2.53 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 168.3, 159.2, 147.9, 141.7, 139.5, 138.3, 136.7, 136.0, 135.8, 134.4, 129.5, 129.1, 127.7, 127.4, 127.1, 121.6, 121.4, 121.1, 116.3, 113.6, 55.0, 19.7; HRMS (ESI) calcd for $\text{C}_{24}\text{H}_{21}\text{N}_2\text{O}_2$ $[\text{M} + \text{H}]^+$ 369.1603, Found 369.1610.

4'-Methyl-3-methyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**3e**)



87% yield (61 mg); colorless oil; ^1H NMR (400 MHz, CDCl_3) δ 9.65 (s, 1H), 8.78 (d, $J = 7.3$ Hz, 1H), 8.61 (d, $J = 4.0$ Hz, 1H), 8.07 (d, $J = 8.2$ Hz, 1H), 7.50 (t, $J = 7.9$ Hz, 1H), 7.46–7.39 (m, 4H), 7.37–7.33 (m, 1H), 7.29–7.24 (m, 2H), 7.01 (d, $J = 7.9$ Hz, 2H), 2.52 (s, 3H), 2.15 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 168.4, 147.9, 139.6, 138.4, 137.4, 136.9, 136.8, 136.0, 135.7, 134.4, 129.2, 129.1, 128.9, 128.5, 127.8, 127.6, 127.2, 121.6, 121.4, 116.5, 20.9, 19.8; HRMS (ESI) calcd for $\text{C}_{24}\text{H}_{21}\text{N}_2\text{O}$ $(\text{M} + \text{H})^+$ 353.1654, found 353.1647.

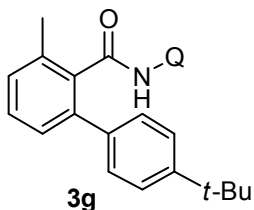
4'-Isopropyl-3-methyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**3f**)



83% yield (63 mg); White solid, mp: 132–134 °C (from ethyl acetate/petroleum ether = 12:1); ^1H NMR (400 MHz, CDCl_3) δ 9.58 (s, 1H), 8.76 (d, $J = 7.6$ Hz, 1H), 8.57 (d, $J = 4.1$ Hz, 1H), 8.03 (dd, $J = 8.2, 1.2$ Hz, 1H), 7.49 (t, $J = 7.9$ Hz, 1H), 7.43–7.41 (m, 3H), 7.39–7.37 (m, 1H), 7.33–7.24 (m, 3H), 7.02 (d, $J = 8.0$ Hz, 2H), 2.68–2.61 (m, 1H), 2.54 (s, 3H), 0.95 (s, 3H), 0.93 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 168.4, 147.8,

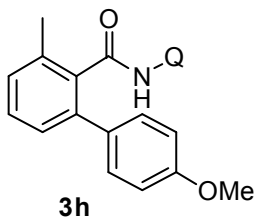
147.7, 139.4, 138.3, 137.7, 136.7, 135.9, 134.4, 129.3, 129.2, 128.5, 127.6,
127.5, 127.2, 126.6, 126.1, 121.5, 121.3, 116.3, 33.5, 24.0, 23.6, 19.8;
HRMS (ESI) calcd for C₂₆H₂₅N₂O (M + H)⁺ 381.1967, found 381.1975.

4'-Tertiary butyl-3-methyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**3g**)



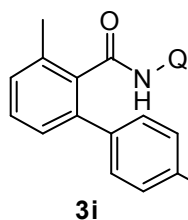
82% yield (65 mg); White solid, mp: 135–137 °C (from ethyl acetate/petroleum ether = 12:1); ¹H NMR (400 MHz, CDCl₃) δ 9.56 (s, 1H), 8.76 (d, *J* = 6.8 Hz, 1H), 8.54 (dd, *J* = 4.1, 1.5 Hz, 1H), 8.01 (d, *J* = 8.2 Hz, 1H), 7.50–7.37 (m, 5H), 7.32–7.25 (m, 3H), 7.16 (d, *J* = 8.3 Hz, 2H), 2.54 (s, 3H), 1.01 (s, 9H); ¹³C NMR (100 MHz, CDCl₃) δ 168.4, 149.9, 147.7, 139.7, 138.2, 137.3, 136.7, 136.0, 135.9, 134.4, 129.3, 129.2, 128.3, 127.6, 127.4, 127.1, 124.9, 121.4, 121.3, 116.2, 34.1, 31.0, 19.8; HRMS (ESI) calcd for C₂₇H₂₇N₂O[M + H]⁺ 395.2123, Found 395.2132.

4'-Methoxy-3-methyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**3h**)



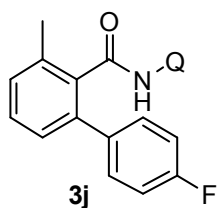
92% yield (68 mg); White solid, mp: 141–143 °C (from ethyl acetate/petroleum ether = 10:1); ¹H NMR (400 MHz, CDCl₃) δ 9.65 (s, 1H), 8.79 (dd, *J* = 7.5, 1.2 Hz, 1H), 8.60 (dd, *J* = 4.2, 1.5 Hz, 1H), 8.04 (dd, *J* = 8.3, 1.5 Hz, 1H), 7.51–7.42 (m, 4H), 7.39–7.31 (m, 2H), 7.27–7.24 (m, 2H), 6.75 (d, *J* = 8.7 Hz, 2H), 3.60 (s, 3H), 2.51 (s, 3H); ¹³C NMR (100 MHz, CDCl₃) δ 168.5, 158.8, 147.9, 139.2, 138.3, 136.7, 136.0, 135.7, 134.4, 132.8, 129.7, 129.1, 129.0, 127.7, 127.5, 127.2, 121.6, 121.4, 116.4, 113.6, 55.0, 19.7; HRMS (ESI) calcd for C₂₄H₂₁N₂O₂ (M + H)⁺ 369.1603, found 369.1600.

4'-Methylthio-3-methyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**3i**)



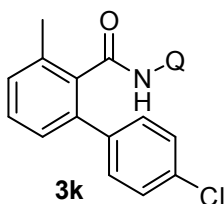
82% yield (63 mg); White solid, mp: 135–137 °C (from ethyl acetate/petroleum ether = 12:1); ^1H NMR (400 MHz, CDCl_3) δ 9.64 (s, 1H), 8.78 (d, J = 7.3 Hz, 1H), 8.60 (d, J = 4.1 Hz, 1H), 8.07 (d, J = 8.3 Hz, 1H), 7.52–7.42 (m, 4H), 7.39–7.34 (m, 2H), 7.29–7.26 (m, 2H), 7.09 (d, J = 7.6 Hz, 2H), 2.52 (s, 3H), 2.28 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 168.3, 148.0, 139.0, 138.4, 137.5, 137.2, 136.7, 136.1, 135.9, 134.3, 129.5, 129.2, 129.0, 127.8, 127.5, 127.2, 126.4, 121.7, 121.5, 116.6, 19.8, 15.6; HRMS (ESI) calcd for $\text{C}_{24}\text{H}_{21}\text{N}_2\text{OS}$ $[\text{M} + \text{H}]^+$ 385.1375, Found 385.1379.

4'-Fluoro-3-methyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**3j**)



83% yield (59 mg); colorless oil; ^1H NMR (400 MHz, CDCl_3) δ 9.64 (s, 1H), 8.77 (dd, J = 7.4, 1.4 Hz, 1H), 8.60 (dd, J = 4.2, 1.6 Hz, 1H), 8.06 (dd, J = 8.3, 1.6 Hz, 1H), 7.51–7.46 (m, 4H), 7.41–7.33 (m, 2H), 7.29–7.24 (m, 2H), 6.90 (t, J = 8.7 Hz, 2H), 2.52 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 168.1, 162.2 (d, J = 244.9 Hz), 148.0, 138.4 (d, J = 19.8 Hz), 136.8, 136.1, 135.8, 134.1, 130.3, 130.2, 129.6, 129.2, 127.7, 127.5, 127.2, 121.8, 121.5, 116.4, 115.2, 114.9, 19.7; HRMS (ESI) calcd for $\text{C}_{23}\text{H}_{18}\text{FN}_2\text{O}$ $[\text{M} + \text{H}]^+$ 357.1403, Found 357.1399.

4'-Chloro-3-methyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**3k**)



82% yield (61 mg); colorless oil; ^1H NMR (400 MHz, CDCl_3) δ 9.66 (s, 1H), 8.77 (d, J = 7.4 Hz, 1H), 8.59 (d, J = 4.0 Hz, 1H), 8.04 (d, J = 8.2 Hz, 1H), 7.49–7.45 (m, 4H), 7.40–7.36 (m, 1H), 7.34–7.31 (m, 1H), 7.29–7.22 (m, 2H), 7.19–7.16 (m, 2H), 2.52 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 168.0, 148.0, 138.8, 138.3, 138.2, 136.7, 136.1, 135.8, 134.1, 133.4, 129.9, 129.7,

129.5, 129.2, 128.3, 127.7, 127.4, 127.1, 121.9, 121.5, 120.1, 116.5, 19.7;

HRMS (ESI) calcd for $C_{23}H_{18}ClN_2O$ ($M + H$)⁺ 373.1108, found 373.1102.

4'-cyano-3-methyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**3l**)

63% yield (46 mg); White solid, mp: 133–135 °C (from ethyl

acetate/petroleum ether = 10:1); ¹H NMR (400 MHz, CDCl₃) δ 9.64 (s, 1H),

8.73 (dd, *J* = 5.9, 3.0 Hz, 1H), 8.61 (dd, *J* = 4.2, 1.5 Hz, 1H), 8.13 (dd, *J* =

8.3, 1.5 Hz, 1H), 7.79–7.77 (m, 2H), 7.70–7.68 (m, 2H), 7.64–7.62 (m, 2H),

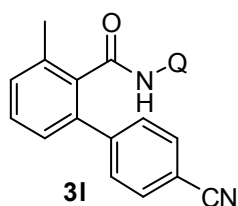
7.53–7.50 (m, 2H), 7.41–7.38 (m, 1H), 7.28–7.26 (m, 1H), 2.54 (s, 3H); ¹³C

NMR (100 MHz, CDCl₃) δ 167.5, 148.1, 145.1, 143.4, 138.2, 137.7, 136.3,

133.9, 132.8, 132.5, 131.9, 130.6, 129.5, 129.3, 127.8, 127.2, 127.1, 122.2,

121.6, 118.4, 116.6, 19.7; HRMS (ESI) calcd for $C_{24}H_{18}N_3O$ [$M + H$]⁺

364.1450, Found 364.1461.



4'-Aldehyde-3-methyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**3m**)

54% yield (40 mg); White solid, mp: 138–140 °C (from ethyl

acetate/petroleum ether = 10:1); ¹H NMR (400 MHz, CDCl₃) δ 9.82 (s, 1H),

9.66 (s, 1H), 8.75 (d, *J* = 7.1 Hz, 1H), 8.59 (d, *J* = 4.1 Hz, 1H), 8.07 (d, *J* =

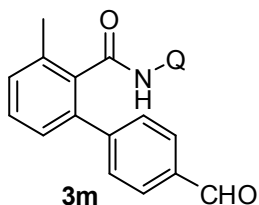
8.3 Hz, 1H), 7.74–7.68 (m, 4H), 7.52–7.42 (m, 3H), 7.36–7.30 (m, 3H), 2.55

(s, 3H); ¹³C NMR (100 MHz, CDCl₃) δ 191.8, 167.7, 148.0, 146.7, 138.3,

136.7, 136.2, 136.1, 135.0, 134.0, 130.4, 129.6, 129.4, 129.3, 128.0, 127.7,

127.3, 127.2, 122.0, 121.5, 116.5, 19.7; HRMS (ESI) calcd for $C_{24}H_{19}N_2O_2$

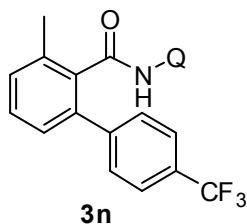
[$M + H$]⁺ 367.1447, Found 367.1450.



4'-Trifluoromethyl-3-methyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**3n**)

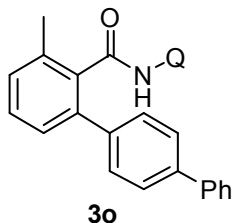
74% yield (60 mg); colorless oil; ¹H NMR (400 MHz, CDCl₃) δ 9.65 (s, 1H),

8.74 (dd, *J* = 7.2, 1.4 Hz, 1H), 8.58 (dd, *J* = 4.1, 1.4 Hz, 1H), 8.07 (dd, *J* =



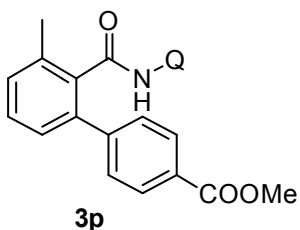
8.3, 1.4 Hz, 1H), 7.64 (d, $J = 8.1$ Hz, 2H), 7.50–7.41 (m, 5H), 7.35–7.33 (m, 2H), 7.28 (d, $J = 7.6$ Hz, 1H), 2.55 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 167.8, 148.0, 144.0, 138.3, 138.1, 136.7, 136.1, 134.0, 130.2, 129.4, 129.0, 127.6 (q, $J = 30.0$ Hz), 127.4, 127.1, 125.1, 125.0, 124.0 (q, $J = 270.4$ Hz), 122.0, 121.5, 118.0, 116.6, 19.8; HRMS (ESI) calcd for $\text{C}_{24}\text{H}_{18}\text{F}_3\text{N}_2\text{O}$ ($\text{M} + \text{H}$) $^+$ 407.1371, found 407.1378.

4'-Phenyl-3-methyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**3o**)



80% yield (66 mg); White solid, mp: 143–145 °C (from ethyl acetate/petroleum ether = 12:1); ^1H NMR (400 MHz, CDCl_3) δ 9.70 (s, 1H), 8.79 (d, $J = 7.4$ Hz, 1H), 8.58 (d, $J = 3.8$ Hz, 1H), 8.01 (d, $J = 7.1$ Hz, 1H), 7.61–7.59 (m, 2H), 7.49–7.35 (m, 7H), 7.32–7.25 (m, 6H), 2.55 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 168.3, 147.9, 140.5, 139.9, 139.3, 139.2, 138.3, 136.7, 136.0, 135.9, 134.3, 129.5, 129.2, 129.0, 128.5, 127.7, 127.6, 127.5, 127.2, 127.1, 126.9, 121.7, 121.4, 116.5, 19.8; HRMS (ESI) calcd for $\text{C}_{29}\text{H}_{23}\text{N}_2\text{O}$ [$\text{M} + \text{H}$] $^+$ 415.1810, Found 415.1803.

Methyl-3'-methyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**3p**)

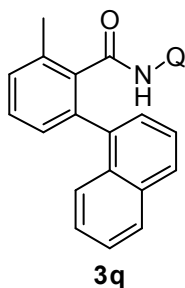


74% yield (57 mg); White solid, mp: 131–133 °C (from ethyl acetate/petroleum ether = 8:1); ^1H NMR (400 MHz, CDCl_3) δ 9.67 (s, 1H), 8.76 (d, $J = 7.4$ Hz, 1H), 8.58 (dd, $J = 4.1, 1.1$ Hz, 1H), 8.03 (dd, $J = 8.2, 1.2$ Hz, 1H), 7.91–7.89 (m, 2H), 7.62–7.60 (m, 2H), 7.50–7.46 (m, 1H), 7.44–7.39 (m, 2H), 7.32–7.28 (m, 3H), 3.76 (s, 3H), 2.53 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 167.8, 166.6, 147.9, 145.0, 138.4, 138.2, 136.6, 136.0, 135.9, 134.0, 130.1, 129.4, 129.2, 128.8, 128.6, 127.7,

127.4, 127.1, 121.8, 121.4, 116.5, 51.8, 19.7; HRMS (ESI) calcd for

$C_{25}H_{21}N_2O_3$ ($M + H$)⁺ 397.1552, found 397.1548.

2-Methyl-N-(quinolin-8-yl)-6-(naph-1-yl)benzamide (**3q**)



47% yield (32 mg); White solid, mp: 143–145 °C (from ethyl

acetate/petroleum ether = 10:1); ¹H NMR (400 MHz, CDCl₃) δ 9.53 (s, 1H),

8.52 (dd, *J* = 7.2, 1.6 Hz, 1H), 8.29 (dd, *J* = 4.1, 1.5 Hz, 1H), 7.93–7.85 (m,

2H), 7.64 (d, *J* = 8.1 Hz, 1H), 7.54–7.36 (m, 6H), 7.32–7.26 (m, 4H),

7.22–7.19 (m, 1H), 2.58 (s, 3H); ¹³C NMR (100 MHz, CDCl₃) δ 167.6,

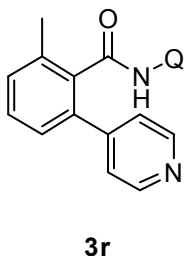
147.4, 138.1, 137.9, 137.8, 137.6, 136.1, 135.8, 134.0, 133.4, 132.2, 129.7,

128.7, 128.6, 128.0, 127.7, 127.4, 127.0, 126.9, 126.3, 125.9, 125.5, 124.9,

121.3, 121.1, 116.0, 19.9; HRMS (ESI) calcd for $C_{27}H_{19}N_2O$ [$M + H$]⁺

387.1497, Found 387.1501.

2-Methyl-N-(quinolin-8-yl)-6-(pyridin-8-yl)benzamide (**3r**)



65% yield (45 mg); colorless oil; ¹H NMR (400 MHz, CDCl₃) δ 9.67 (s, 1H),

8.76 (dd, *J* = 6.9, 2.0 Hz, 1H), 8.62 (d, *J* = 4.2 Hz, 1H), 8.44 (d, *J* = 5.2 Hz,

2H), 8.11 (dd, *J* = 8.3, 1.5 Hz, 1H), 7.55–7.50 (m, 2H), 7.48–7.45 (m, 3H),

7.39–7.36 (m, 2H), 7.31–7.29 (m, 1H), 2.55 (s, 3H); ¹³C NMR (100 MHz,

CDCl₃) δ 167.5, 149.5, 148.3, 148.1, 138.3, 136.8, 136.6, 136.2, 134.0,

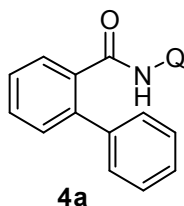
130.8, 129.5, 127.8, 127.2, 127.1, 123.5, 122.1, 121.6, 116.6, 19.7; HRMS

(ESI) calcd for $C_{22}H_{16}N_3O$ [$M + H$]⁺ 338.1293, Found 338.1294.

N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**4a**)

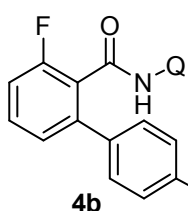
91% yield (59 mg); colorless oil; ¹H NMR (400 MHz, CDCl₃) δ 9.79 (s, 1H),

8.81 (d, *J* = 7.4 Hz, 1H), 8.51 (d, *J* = 2.6 Hz, 1H), 8.05 (d, *J* = 8.2 Hz, 1H),



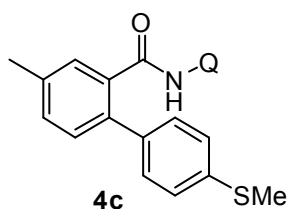
7.91 (d, $J = 7.3$ Hz, 1H), 7.57–7.45 (m, 6H), 7.37–7.33 (m, 1H), 7.30–7.26 (m, 2H), 7.18–7.16 (m, 2H); ^{13}C NMR (100 MHz, CDCl_3) δ 167.8, 147.7, 140.5, 140.3, 138.4, 135.9, 134.5, 130.7, 130.5, 129.4, 129.1, 128.9, 128.7, 128.3, 128.1, 127.6, 127.3, 121.5, 121.4, 116.2; HRMS (ESI) calcd for $\text{C}_{22}\text{H}_{17}\text{N}_2\text{O}$ ($\text{M} + \text{H}$) $^+$ 325.1341, found 325.1355.

4'-Methoxy-3-fluoro-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**4b**)



67% yield (50 mg); White solid, mp: 128–130 °C (from ethyl acetate/petroleum ether = 10:1); ^1H NMR (400 MHz, CDCl_3) δ 9.89 (s, 1H), 8.80 (d, $J = 6.8$ Hz, 1H), 8.66 (d, $J = 4.2$ Hz, 1H), 8.11 (d, $J = 8.3$ Hz, 1H), 7.53–7.51 (m, 1H), 7.48–7.40 (m, 3H), 7.39–7.38 (m, 1H), 7.27–7.24 (m, 2H), 7.19–7.14 (m, 1H), 6.80 (d, $J = 8.3$ Hz, 2H), 3.68 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 163.5, 160.6 (d, $J = 249.8$ Hz), 148.1, 142.0, 138.4, 136.2, 134.5, 134.2, 131.4, 130.9, 130.8, 129.7, 127.8, 127.3, 125.8, 121.9, 121.5, 116.7, 114.4 (d, $J = 21.7$ Hz), 113.9, 55.1; HRMS (ESI) calcd for $\text{C}_{23}\text{H}_{18}\text{FN}_2\text{O}_2$ ($\text{M} + \text{H}$) $^+$ 373.1352, found 373.1350.

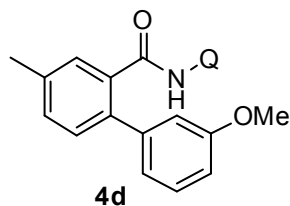
4'-Methylthio-4-methyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**4c**)



72% yield (55 mg); White solid, mp: 127–129 °C (from ethyl acetate/petroleum ether = 12:1); ^1H NMR (400 MHz, CDCl_3) δ 9.78 (s, 1H), 8.81 (dd, $J = 7.6, 1.1$ Hz, 1H), 8.51 (dd, $J = 4.2, 1.6$ Hz, 1H), 8.04 (dd, $J = 8.3, 1.6$ Hz, 1H), 7.72 (s, 1H), 7.52–7.48 (m, 1H), 7.44–7.39 (m, 3H), 7.34–7.30 (m, 3H), 7.14–7.12 (m, 2H), 2.45 (s, 3H), 2.26 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 167.9, 147.7, 138.3, 137.7, 137.4, 136.6,

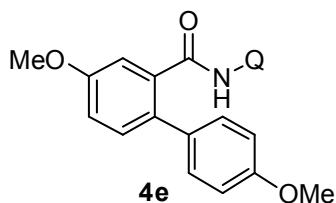
135.9, 135.6, 134.5, 131.3, 130.4, 129.8, 129.3, 127.6, 127.1, 126.5, 121.5, 121.3, 116.2, 21.0, 15.6; HRMS (ESI) calcd for C₂₄H₂₁N₂OS (M + H)⁺ 385.1375, found 385.1370.

3'-Methoxy-4-chloro-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**4d**)



79% yield (58 mg); White solid, mp: 132–134 °C (from ethyl acetate/petroleum ether = 10:1); ¹H NMR (400 MHz, CDCl₃) δ 9.77 (s, 1H), 8.82 (d, *J* = 7.6 Hz, 1H), 8.48 (dd, *J* = 4.2, 1.5 Hz, 1H), 7.97 (dd, *J* = 8.3, 1.4 Hz, 1H), 7.71 (s, 1H), 7.47–7.43 (m, 1H), 7.38–7.30 (m, 3H), 7.27–7.24 (m, 1H), 7.12–7.08 (m, 1H), 7.06–7.04 (m, 2H), 6.64 (d, *J* = 7.1 Hz, 1H), 3.63 (s, 3H), 2.43 (s, 3H); ¹³C NMR (100 MHz, CDCl₃) δ 167.9, 159.5, 147.6, 141.2, 138.2, 137.5, 137.1, 135.8, 134.5, 131.1, 130.3, 129.6, 129.2, 127.5, 127.0, 121.3, 121.3, 116.0, 114.0, 113.3, 55.0, 20.9; HRMS (ESI) calcd for C₂₄H₂₁N₂O₂ (M + H)⁺ 369.1603, found 369.1609.

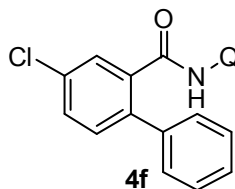
4,4'-dimethoxy-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**4e**)



84% yield (65 mg); White solid, mp: 141–143 °C (from ethyl acetate/petroleum ether = 8:1); ¹H NMR (400 MHz, CDCl₃) δ 9.79 (s, 1H), 8.82 (d, *J* = 7.5 Hz, 1H), 8.50 (d, *J* = 3.8 Hz, 1H), 8.07 (d, *J* = 8.2 Hz, 1H), 7.54–7.50 (m, 1H), 7.46–7.44 (m, 2H), 7.41–7.33 (m, 4H), 7.10–7.07 (m, 1H), 6.79 (d, *J* = 8.4 Hz, 2H), 3.90 (s, 3H), 3.63 (s, 3H); ¹³C NMR (100 MHz, CDCl₃) δ 167.7, 159.1, 158.8, 147.7, 138.5, 136.7, 135.9, 134.5, 132.4, 132.1, 131.9, 130.2, 127.7, 127.2, 121.5,

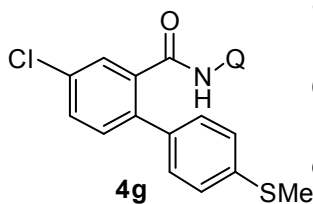
121.3, 117.1, 116.3, 113.9, 113.7, 55.6, 55.1; HRMS (ESI) calcd for $C_{24}H_{21}N_2O_3$ ($M + H$)⁺ 385.1552, found 385.1546.

4-Chloro-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**4f**)



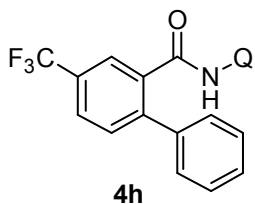
55% yield (39 mg); colorless oil; ¹H NMR (400 MHz, CDCl₃) δ 9.56 (s, 1H), 8.63 (d, *J* = 3.0 Hz, 1H), 8.35 (d, *J* = 7.2 Hz, 1H), 8.05 (d, *J* = 8.3 Hz, 1H), 7.65–7.63 (m, 1H), 7.53–7.51 (m, 2H), 7.45–7.41 (m, 3H), 7.38–7.34 (m, 1H), 7.26–7.22 (m, 3H), 7.17–7.15 (m, 1H); ¹³C NMR (100 MHz, CDCl₃) δ 166.0, 147.9, 139.3, 138.7, 138.7, 136.8, 136.0, 130.5, 130.3, 129.5, 129.4, 128.6, 128.3, 127.9, 127.8, 127.6, 127.1, 121.5, 121.4, 116.4; HRMS (ESI) calcd for $C_{22}H_{16}ClN_2O$ ($M + H$)⁺ 359.0951, found 359.0947.

4'-Methylthio-4-chloro-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**4g**)



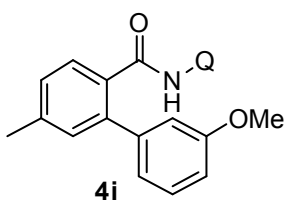
71% yield (57 mg); White solid, mp: 121–123 °C (from ethyl acetate/petroleum ether = 12:1); ¹H NMR (400 MHz, CDCl₃) δ 9.80 (s, 1H), 8.77 (d, *J* = 7.3 Hz, 1H), 8.53 (d, *J* = 4.1 Hz, 1H), 8.08 (d, *J* = 8.3 Hz, 1H), 7.90 (d, *J* = 4.0 Hz, 1H), 7.52–7.47 (m, 3H), 7.40–7.37 (m, 4H), 7.16–7.14 (m, 2H), 2.29 (s, 3H); ¹³C NMR (100 MHz, CDCl₃) δ 166.2, 147.8, 138.7, 138.3, 137.9, 137.1, 136.0, 135.4, 134.2, 133.7, 131.8, 130.6, 129.3, 129.3, 127.7, 127.2, 126.5, 121.8, 121.5, 116.5, 15.5; HRMS (ESI) calcd for $C_{23}H_{18}ClN_2OS$ ($M + H$)⁺ 405.0828, found 405.0834.

4-Trifluoro-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**4h**)



77% yield (60 mg); White solid, mp: 123–125 °C (from ethyl acetate/petroleum ether = 12:1); ^1H NMR (400 MHz, CDCl_3) δ 9.78 (s, 1H), 8.78 (dd, $J = 7.4, 1.2$ Hz, 1H), 8.49 (dd, $J = 4.2, 1.6$ Hz, 1H), 8.20 (s, 1H), 8.05 (dd, $J = 8.3, 1.5$ Hz, 1H), 7.79 (d, $J = 8.0$ Hz, 1H), 7.59 (d, $J = 8.2$ Hz, 1H), 7.52–7.46 (m, 4H), 7.33–7.28 (m, 3H), 7.21–7.17 (m, 1H); ^{13}C NMR (100 MHz, CDCl_3) δ 166.2, 147.8, 143.5, 138.5, 138.2, 136.5, 135.9, 134.0, 131.2, 130.2, 128.8, 128.6, 128.4, 127.5 (q, $J = 24.1$ Hz), 127.1, 127.0, 126.5, 121.9, 121.5, 118.9 (q, $J = 276.2$ Hz), 116.4; HRMS (ESI) calcd for $\text{C}_{23}\text{H}_{16}\text{F}_3\text{N}_2\text{O}$ ($\text{M} + \text{H}$) $^+$ 393.1215, found 393.1208.

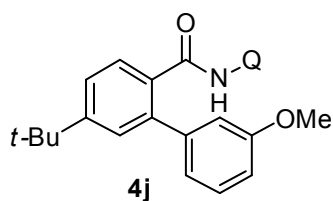
3'-Methoxy-5-methyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**4i**)



73% yield (54 mg); White solid, mp: 131–133 °C (from ethyl acetate/petroleum ether = 10:1); ^1H NMR (400 MHz, CDCl_3) δ 9.77 (s, 1H), 8.81 (d, $J = 7.5$ Hz, 1H), 8.49 (dd, $J = 4.1, 1.5$ Hz, 1H), 8.00 (dd, $J = 8.3, 1.4$ Hz, 1H), 7.83 (d, $J = 8.4$ Hz, 1H), 7.47 (t, $J = 7.9$ Hz, 1H), 7.40–7.38 (m, 1H), 7.30–7.27 (m, 3H), 7.15–7.11 (m, 1H), 7.07–7.05 (m, 2H), 6.67 (d, $J = 8.8$ Hz, 1H), 3.67 (s, 3H), 2.43 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 167.7, 159.5, 147.6, 141.5, 140.6, 140.1, 138.3, 135.8, 134.6, 133.2, 131.1, 129.3, 129.2, 128.3, 127.6, 127.1, 121.4, 121.3, 116.0, 114.0, 113.6, 55.1, 21.3; HRMS (ESI) calcd for $\text{C}_{24}\text{H}_{21}\text{N}_2\text{O}_2$ ($\text{M} + \text{H}$) $^+$ 369.1603, found 369.1601.

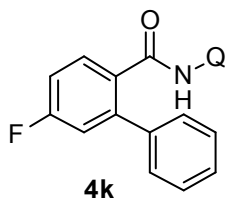
3'-Methoxy-5-tertbutyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**4j**)

71% yield (58 mg); White solid, mp: 134–136 °C (from ethyl



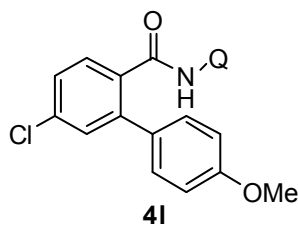
acetate/petroleum ether = 10:1); ^1H NMR (400 MHz, CDCl_3) δ 9.79 (s, 1H), 8.82 (d, J = 7.6 Hz, 1H), 8.51 (dd, J = 4.1, 1.5 Hz, 1H), 8.04 (dd, J = 8.3, 1.4 Hz, 1H), 7.87 (d, J = 8.1 Hz, 1H), 7.54–7.47 (m, 3H), 7.42–7.41 (m, 1H), 7.33–7.30 (m, 1H), 7.18–7.14 (m, 1H), 7.10–7.08 (m, 2H), 6.70 (d, J = 7.2 Hz, 1H), 3.69 (s, 3H), 1.39 (s, 9H); ^{13}C NMR (100 MHz, CDCl_3) δ 167.8, 159.5, 153.8, 147.6, 141.9, 139.8, 138.4, 135.9, 134.6, 133.3, 129.3, 129.1, 127.6, 127.5, 127.2, 124.8, 121.5, 121.3, 116.1, 114.3, 113.4, 55.1, 34.9, 31.2; HRMS (ESI) calcd for $\text{C}_{27}\text{H}_{27}\text{N}_2\text{O}_2$ ($\text{M} + \text{H}$) $^+$ 411.2073, found 411.2069.

5-Fluoro-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**4k**)



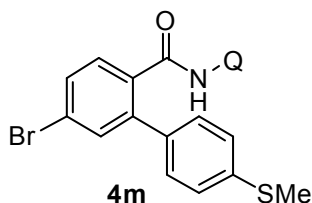
81% yield (55 mg); White solid, mp: 123–125 °C (from ethyl acetate/petroleum ether = 12:1); ^1H NMR (400 MHz, CDCl_3) δ 9.60 (s, 1H), 8.55 (dd, J = 4.2, 1.6 Hz, 1H), 8.49 (dd, J = 8.1, 1.4 Hz, 1H), 8.00 (dd, J = 8.3, 1.6 Hz, 1H), 7.54–7.51 (m, 3H), 7.38–7.37 (m, 1H), 7.36 (s, 1H), 7.31–7.28 (m, 1H), 7.25–7.21 (m, 3H), 7.17–7.15 (m, 2H); ^{13}C NMR (100 MHz, CDCl_3) δ 166.7, 162.2 (d, J = 248.4 Hz), 147.8, 143.2, 143.1, 139.3, 138.1, 135.9, 134.1, 132.5, 128.8, 128.4, 128.3, 127.8, 127.5, 127.1, 121.6, 121.4 (d, J = 18.9 Hz), 116.3, 116.0 (d, J = 21.6 Hz); HRMS (ESI) calcd for $\text{C}_{22}\text{H}_{16}\text{FN}_2\text{O}$ ($\text{M} + \text{H}$) $^+$ 343.1247, found 343.1250.

4'-Methoxy-5-chloro-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**4l**)



57% yield (44 mg); White solid, mp: 124–126 °C (from ethyl acetate/petroleum ether = 10:1); ^1H NMR (400 MHz, CDCl_3) δ 9.60 (s, 1H), 8.58 (dd, J = 4.2, 1.6 Hz, 1H), 8.52 (d, J = 8.1 Hz, 1H), 8.04 (dd, J = 8.3, 1.6 Hz, 1H), 7.45–7.40 (m, 7H), 7.35–7.32 (m, 1H), 6.94 (d, J = 8 Hz, 2H), 3.65 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 167.0, 159.2, 149.2, 147.9, 141.9, 135.9, 134.7, 134.4, 134.1, 131.5, 129.7, 128.7, 127.5, 127.1, 126.8, 124.1, 121.4, 116.4, 114.2, 113.8, 55.1; HRMS (ESI) calcd for $\text{C}_{23}\text{H}_{18}\text{ClN}_2\text{O}_2$ ($\text{M} + \text{H}$) $^+$ 389.1057, found 389.1048.

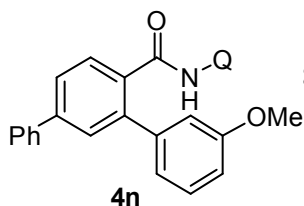
4'-Methylthio-5-bromo-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**4m**)



70% yield (63 mg); White solid, mp: 127–129 °C (from ethyl acetate/petroleum ether = 12:1); ^1H NMR (400 MHz, CDCl_3) δ 9.87 (s, 1H), 8.83 (d, J = 7.6 Hz, 1H), 8.55 (dd, J = 4.1, 1.5 Hz, 1H), 8.09 (dd, J = 8.3, 1.4 Hz, 1H), 8.02 (d, J = 8.0 Hz, 1H), 7.68–7.66 (m, 2H), 7.50–7.46 (m, 4H), 7.37–7.34 (m, 1H), 7.20–7.18 (m, 2H), 2.30 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 167.4, 147.8, 143.5, 139.9, 138.3, 136.7, 135.9, 134.5, 130.1, 129.5, 129.3, 128.9, 127.9, 127.7, 127.2, 126.6, 126.2, 121.5, 121.4, 116.3, 15.6; HRMS (ESI) calcd for $\text{C}_{23}\text{H}_{18}\text{BrN}_2\text{OS}$ ($\text{M} + \text{H}$) $^+$ 449.0323, found 449.0320.

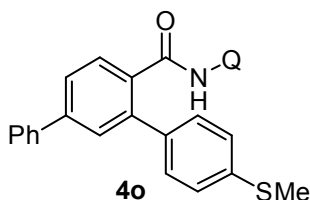
3'-Methoxy-5-Phenyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (**4n**)

72% yield (62 mg); White solid, mp: 144–146 °C (from ethyl acetate/petroleum ether = 10:1); ^1H NMR (400 MHz, CDCl_3) δ 9.84 (s, 1H), 8.83 (d, J = 7.5 Hz, 1H), 8.54 (dd, J = 4.2, 1.6 Hz, 1H), 8.08 (dd, J



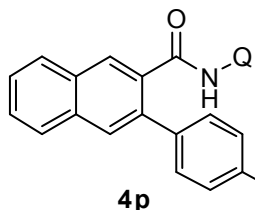
= 8.2, 1.4 Hz, 1H), 8.01 (d, J = 7.8 Hz, 1H), 7.74–7.67 (m, 4H), 7.52–7.47 (m, 5H), 7.37–7.34 (m, 1H), 7.20–7.12 (m, 3H), 6.73 (d, J = 8.2 Hz, 1H), 3.71 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 167.5, 159.7, 147.7, 143.4, 141.4, 140.7, 140.0, 138.4, 135.9, 134.8, 134.6, 129.9, 129.4, 129.3, 128.9, 127.9, 127.7, 127.3, 126.3, 121.5, 121.5, 121.4, 116.2, 114.2, 113.9, 55.2; HRMS (ESI) calcd for $\text{C}_{29}\text{H}_{23}\text{N}_2\text{O}_2$ ($\text{M} + \text{H}$) $^+$ 431.1760, found 431.1755.

4'-Methylthio-5-Phenyl-N-(quinolin-8-yl)-[1,1'-biphenyl]-2-carboxamide (4o)



71% yield (63 mg); White solid, mp: 142–144 °C (from ethyl acetate/petroleum ether = 12:1); ^1H NMR (400 MHz, CDCl_3) δ 9.69 (s, 1H), 8.55 (dd, J = 4.0, 1.3 Hz, 2H), 8.02 (dd, J = 8.3, 1.5 Hz, 1H), 7.68–7.65 (m, 4H), 7.52–7.46 (m, 6H), 7.33–7.30 (m, 1H), 7.14–7.12 (m, 4H), 2.30 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 167.5, 147.8, 142.3, 140.6, 139.9, 138.3, 137.9, 136.9, 135.9, 134.6, 134.1, 130.3, 129.0, 128.9, 128.0, 127.9, 127.6, 127.2, 127.1, 126.3, 121.7, 121.4, 116.5, 116.1, 15.5; HRMS (ESI) calcd for $\text{C}_{29}\text{H}_{23}\text{N}_2\text{OS}$ ($\text{M} + \text{H}$) $^+$ 447.1531, found 447.1526.

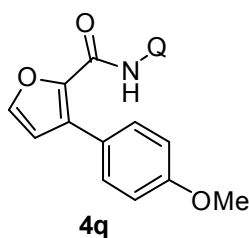
3-(4-methoxyphenyl)-N-(quinolin-8-yl)-2-naphthamide (4p)



73% yield (59 mg); White solid, mp: 147–149 °C (from ethyl acetate/petroleum ether = 8:1); ^1H NMR (400 MHz, CDCl_3) δ 9.98 (s, 1H), 8.86 (d, J = 7.5 Hz, 1H), 8.57 (d, J = 4.0 Hz, 1H), 8.43 (s, 1H), 8.10 (d, J = 8.0 Hz, 1H), 7.97 (d, J = 8.0 Hz, 1H), 7.91–7.89 (m, 2H),

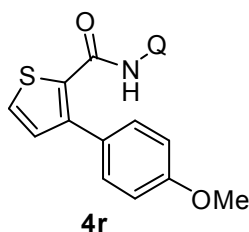
7.59–7.53 (m, 4H), 7.50–7.46 (m, 2H), 7.37 (dd, $J = 8.2, 4.2$ Hz, 1H), 6.86 (d, $J = 8.4$ Hz, 2H), 3.69 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 167.8, 159.3, 147.8, 138.5, 137.0, 136.0, 134.7, 134.4, 134.1, 132.6, 131.8, 131.6, 130.3, 129.6, 129.6, 128.5, 127.7, 127.7, 127.3, 126.6, 121.5, 121.4, 116.4, 114.0, 55.2; HRMS (ESI) calcd for $\text{C}_{27}\text{H}_{21}\text{N}_2\text{O}_2$ ($\text{M} + \text{H}$) $^+$ 405.1603, found 405.1611.

2-(4-methoxyphenyl)-N-(quinolin-8-yl)furan-3-carboxamide (**4q**)



71% yield (49 mg); White solid, mp: 153–155 °C (from ethyl acetate/petroleum ether = 12:1); ^1H NMR (400 MHz, CDCl_3) δ 10.79 (s, 1H), 8.87 (d, $J = 6.9$ Hz, 1H), 8.83 (d, $J = 4.1$ Hz, 1H), 8.16 (d, $J = 8.2$ Hz, 1H), 7.74 (d, $J = 8.5$ Hz, 2H), 7.63 (s, 1H), 7.54–7.52 (m, 2H), 7.46 (dd, $J = 8.3, 4.2$ Hz, 1H), 6.99 (d, $J = 8.5$ Hz, 2H), 6.68 (s, 1H), 3.86 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 159.7, 157.0, 148.2, 143.3, 141.6, 138.8, 136.3, 134.5, 132.1, 130.8, 128.0, 127.4, 124.1, 121.6, 121.5, 116.7, 114.8, 113.7, 55.3; HRMS (ESI) calcd for $\text{C}_{21}\text{H}_{17}\text{N}_2\text{O}_3$ ($\text{M} + \text{H}$) $^+$ 345.1239, found 345.1245.

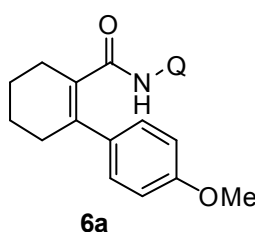
2-(4-methoxyphenyl)-N-(quinolin-8-yl)thiophene-3-carboxamide (**4r**)



70% yield (50 mg); White solid, mp: 137–139 °C (from ethyl acetate/petroleum ether = 10:1); ^1H NMR (400 MHz, CDCl_3) δ 10.14 (s, 1H), 8.83 (d, $J = 7.6$ Hz, 1H), 8.31 (d, $J = 3.8$ Hz, 1H), 8.04 (d, $J = 8.2$ Hz, 1H), 7.53–7.42 (m, 5H), 7.31 (dd, $J = 8.2, 4.1$ Hz, 1H), 7.08 (d, $J = 5.0$ Hz, 1H), 7.00–6.98 (m, 2H), 3.83 (s, 3H); ^{13}C NMR (100 MHz,

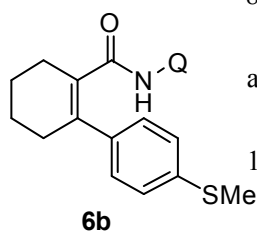
CDCl₃) δ 160.8, 160.1, 147.4, 142.9, 138.4, 135.8, 135.4, 134.6, 131.5, 130.8, 129.2, 127.7, 127.4, 127.3, 121.4, 121.3, 116.4, 114.6, 55.3; HRMS (ESI) calcd for C₂₁H₁₇N₂O₂S (M + H)⁺ 361.1011, found 361.1006.

4'-methoxy-N-(quinolin-8-yl)-3,4,5,6-tetrahydro-[1,1'-biphenyl]-2-carboxamide (**6a**)



77% yield (55 mg); White solid, mp: 123–125 °C (from ethyl acetate/petroleum ether = 12:1); ¹H NMR (400 MHz, CDCl₃) δ 9.37 (s, 1H), 8.70 (d, *J* = 7.5 Hz, 1H), 8.52 (d, *J* = 3.9 Hz, 1H), 8.02 (d, *J* = 8.2 Hz, 1H), 7.47–7.43 (m, 1H), 7.38 (d, *J* = 8.1 Hz, 1H), 7.31 (dd, *J* = 8.2, 4.2 Hz, 1H), 7.27–7.26 (m, 2H), 6.68 (d, *J* = 8.5 Hz, 2H), 3.54 (s, 3H), 2.63–2.58 (m, 2H), 2.50–2.45 (m, 2H), 1.84–1.76 (m, 4H); ¹³C NMR (100 MHz, CDCl₃) δ 169.9, 159.0, 147.5, 140.2, 138.4, 135.9, 134.7, 134.2, 132.5, 128.8, 127.7, 127.2, 121.2, 121.0, 116.0, 113.7, 55.0, 32.0, 27.2, 22.8, 22.2; HRMS (ESI) calcd for C₂₃H₂₃N₂O₂ (M + H)⁺ 359.1760, found 359.1766.

4'-(methylthio)-N-(quinolin-8-yl)-3,4,5,6-tetrahydro-[1,1'-biphenyl]-2-carboxamide (**6b**)

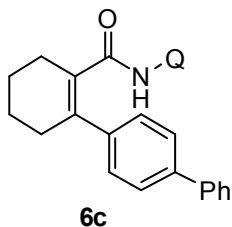


81% yield (61 mg); White solid, mp: 113–115 °C (from ethyl acetate/petroleum ether = 12:1); ¹H NMR (400 MHz, CDCl₃) δ 9.37 (s, 1H), 8.68 (d, *J* = 7.5 Hz, 1H), 8.52 (d, *J* = 4.1 Hz, 1H), 8.03 (d, *J* = 8.2 Hz, 1H), 7.47–7.43 (m, 1H), 7.40–7.38 (m, 1H), 7.32 (dd, *J* = 8.2, 4.2 Hz, 1H), 7.27–7.23 (m, 2H), 7.03 (d, *J* = 8.1 Hz, 2H), 2.64–2.59 (m, 2H), 2.49–2.45 (m, 2H), 2.18 (s, 3H), 1.83–1.78 (m, 4H); ¹³C NMR (100 MHz, CDCl₃) δ 169.6, 147.6, 139.9, 138.6, 138.4, 137.5, 135.9, 134.5, 133.1, 128.1, 127.6,

127.2, 126.6, 121.2, 121.1, 116.1, 31.7, 27.2, 22.7, 22.1, 15.7; HRMS (ESI)

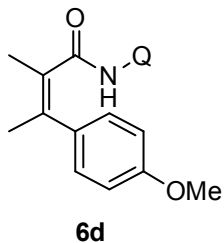
calcd for $C_{23}H_{23}N_2OS$ ($M + H$)⁺ 375.1531, found 375.1526.

N-(quinolin-8-yl)-3,4,5,6-tetrahydro-[1,1':4',1''-terphenyl]-2-carboxamide (**6c**)



63% yield (51 mg); White solid, mp: 144–146 °C (from ethyl acetate/petroleum ether = 12:1); ¹H NMR (400 MHz, CDCl₃) δ 9.41 (s, 1H), 8.70 (d, *J* = 7.6 Hz, 1H), 8.45 (d, *J* = 3.9 Hz, 1H), 7.97 (d, *J* = 8.2 Hz, 1H), 7.47–7.34 (m, 6H), 7.31–7.20 (m, 6H), 2.70–2.60 (m, 2H), 2.59–2.52 (m, 2H), 1.89–1.80 (m, 4H); ¹³C NMR (100 MHz, CDCl₃) δ 169.6, 147.5, 140.7, 140.6, 140.2, 140.2, 138.4, 135.8, 134.6, 133.2, 128.5, 128.1, 127.6, 127.2, 127.1, 126.9, 126.8, 121.2, 121.1, 116.1, 31.8, 27.2, 22.7, 22.1; HRMS (ESI) calcd for $C_{28}H_{25}N_2O$ ($M + H$)⁺ 405.1967, found 405.1957.

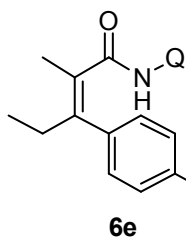
(Z)-3-(4-methoxyphenyl)-2-methyl-N-(quinolin-8-yl)but-2-enamide (**6d**)



63% yield (42 mg); colorless oil; ¹H NMR (400 MHz, CDCl₃) δ 9.99 (s, 1H), 8.90 (d, *J* = 7.4 Hz, 1H), 8.83 (d, *J* = 4.2 Hz, 1H), 8.19 (d, *J* = 8.3 Hz, 1H), 7.59–7.55 (m, 2H), 7.47 (dd, *J* = 8.2, 4.2 Hz, 1H), 7.26–7.22 (m, 2H), 6.94 (d, *J* = 8.4 Hz, 2H), 3.85 (s, 3H), 2.23 (s, 3H), 1.99 (s, 3H); ¹³C NMR (100 MHz, CDCl₃) δ 170.6, 158.6, 148.3, 138.6, 138.0, 137.6, 136.4, 134.7, 134.6, 129.6, 129.0, 128.0, 127.5, 121.6, 116.6, 113.7, 55.3, 22.7, 17.9; HRMS (ESI) calcd for $C_{21}H_{21}N_2O_2$ ($M + H$)⁺ 333.1603, found 333.1600.

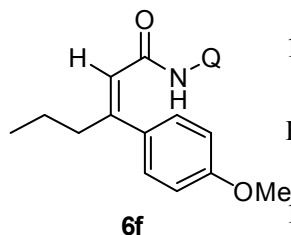
(Z)-3-(4-methoxyphenyl)-2-methyl-N-(quinolin-8-yl)pent-2-enamide (**6e**)

61% yield (42 mg); colorless oil; ¹H NMR (400 MHz, CDCl₃) δ 9.98 (s,



1H), 8.91 (d, $J = 7.3$ Hz, 1H), 8.83 (d, $J = 4.0$ Hz, 1H), 8.19 (d, $J = 8.2$ Hz, 1H), 7.61–7.53 (m, 2H), 7.47 (dd, $J = 8.2, 4.1$ Hz, 1H), 7.18 (d, $J = 8.2$ Hz, 2H), 6.94 (d, $J = 8.3$ Hz, 2H), 3.85 (s, 3H), 2.58 (q, $J = 7.2$ Hz, 2H), 1.93 (s, 3H), 0.99 (t, $J = 7.4$ Hz, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 170.7, 158.6, 148.2, 143.8, 138.5, 136.4, 134.7, 132.8, 129.6, 129.3, 128.0, 127.5, 121.6, 121.6, 116.6, 113.6, 55.2, 29.2, 17.9, 13.3; HRMS (ESI) calcd for $\text{C}_{22}\text{H}_{23}\text{N}_2\text{O}_2$ ($\text{M} + \text{H}$) $^+$ 347.1760, found 347.1751.

(Z)-3-(4-methoxyphenyl)-N-(quinolin-8-yl)hex-2-enamide (6f)



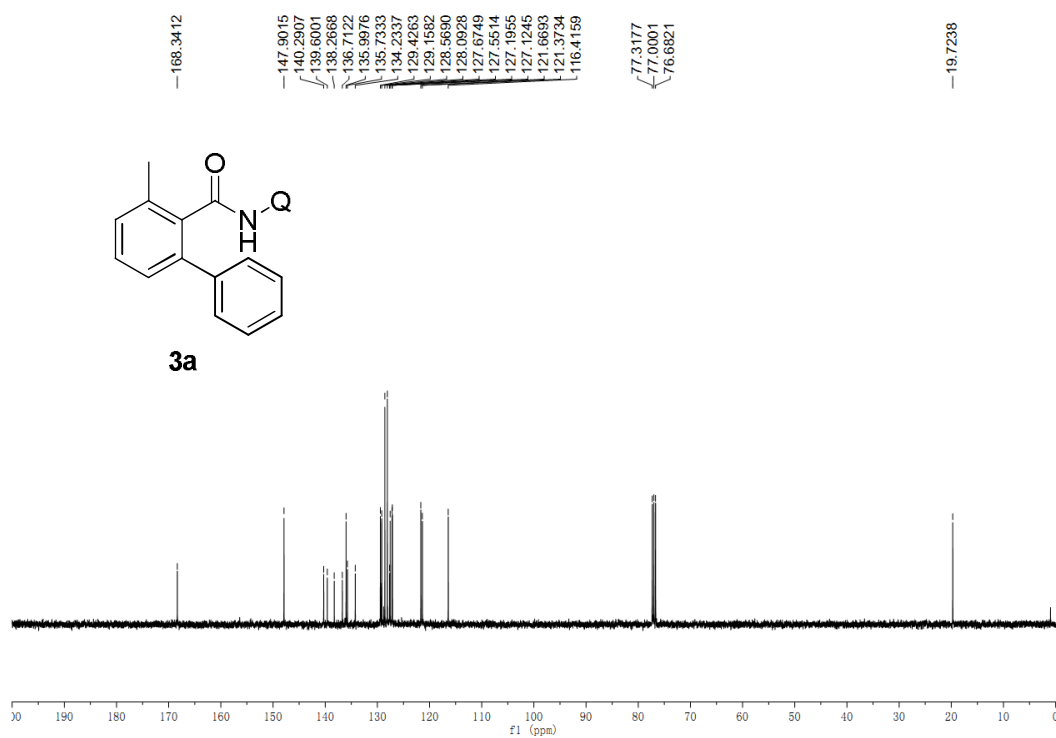
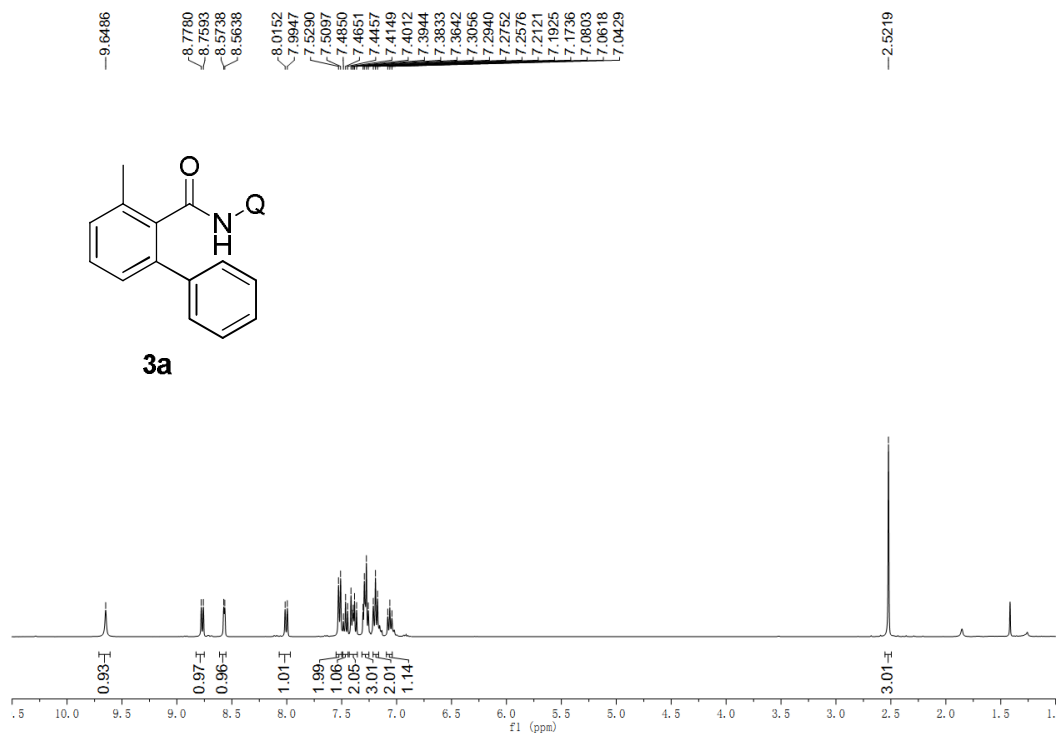
65% yield (45 mg); colorless oil; ^1H NMR (400 MHz, CDCl_3) δ 9.60 (s, 1H), 8.74 (d, $J = 7.4$ Hz, 1H), 8.51 (d, $J = 3.6$ Hz, 1H), 8.06 (d, $J = 8.2$ Hz, 1H), 7.48–7.41 (m, 3H), 7.34 (dd, $J = 8.1, 4.2$ Hz, 1H), 7.28–7.26 (m, 1H), 6.86 (d, $J = 8.3$ Hz, 2H), 6.11 (s, 1H), 3.72 (s, 3H), 2.50 (t, $J = 7.4$ Hz, 2H), 1.46 (q, $J = 7.4$ Hz, 2H), 0.95 (t, $J = 7.3$ Hz, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 165.2, 159.7, 153.6, 147.5, 138.4, 135.9, 134.8, 131.3, 129.2, 127.8, 127.4, 122.3, 121.3, 121.0, 116.3, 114.0, 55.1, 42.4, 20.8, 13.6; HRMS (ESI) calcd for $\text{C}_{22}\text{H}_{23}\text{N}_2\text{O}_2$ ($\text{M} + \text{H}$) $^+$ 347.1760, found 347.1755.

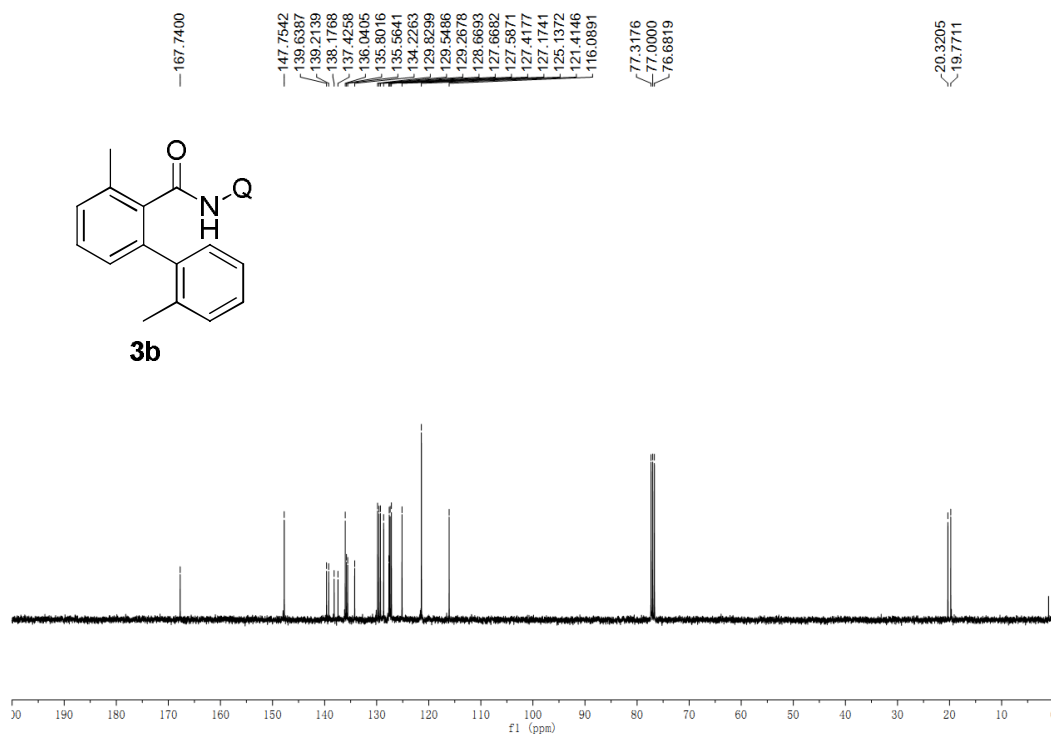
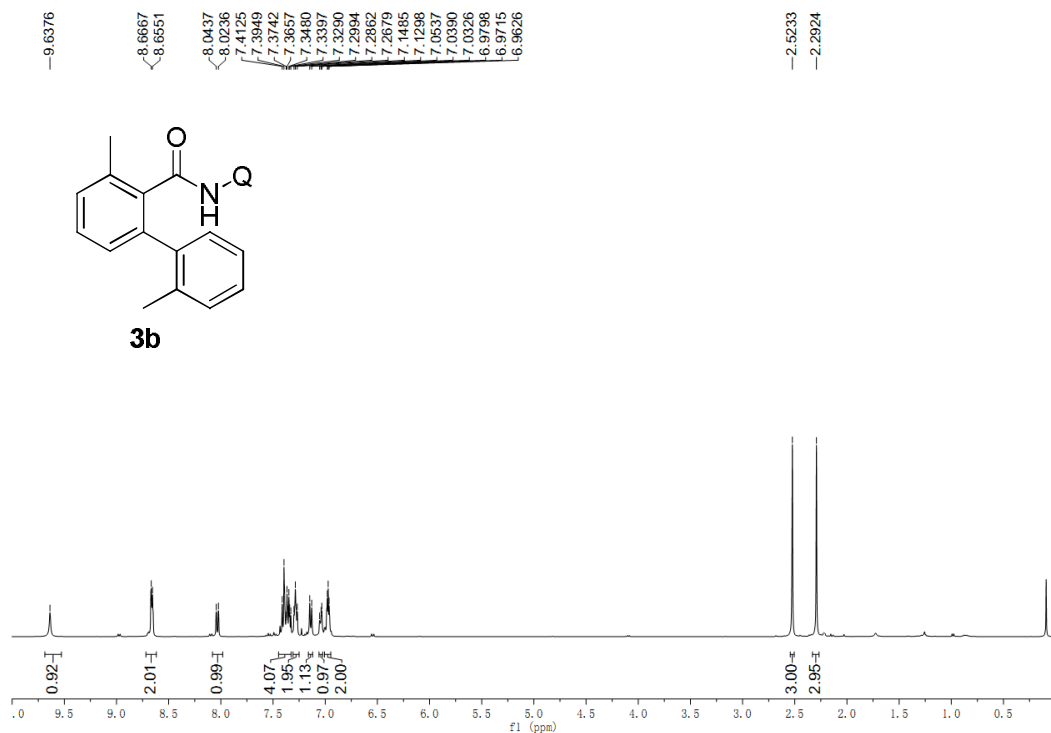
References

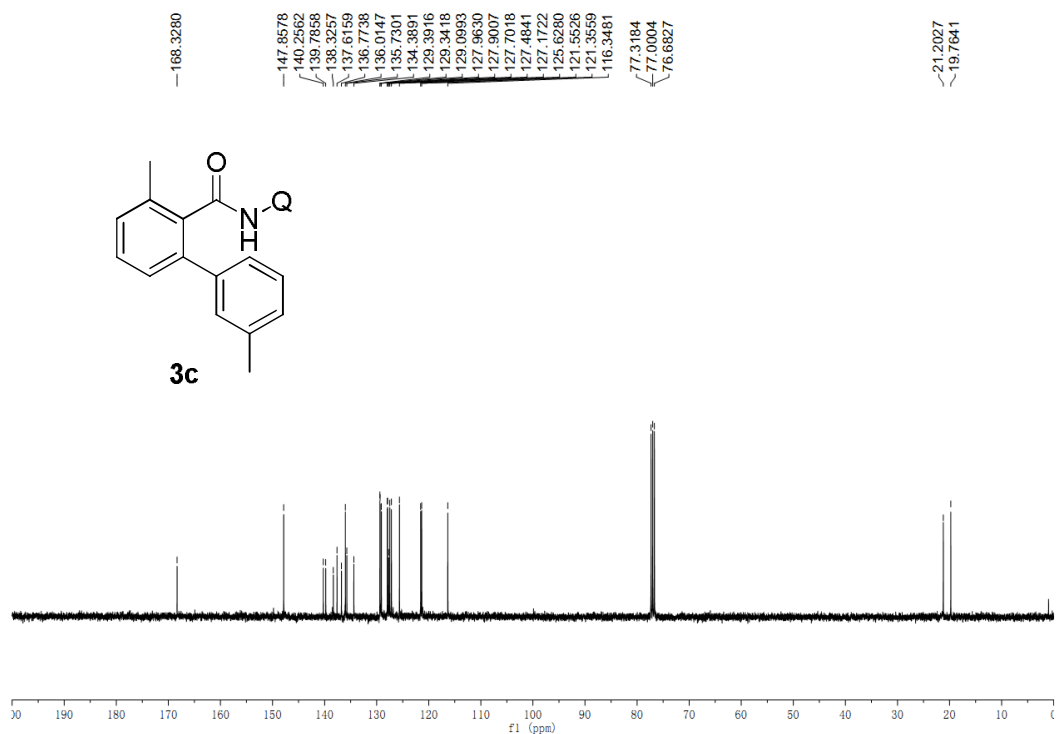
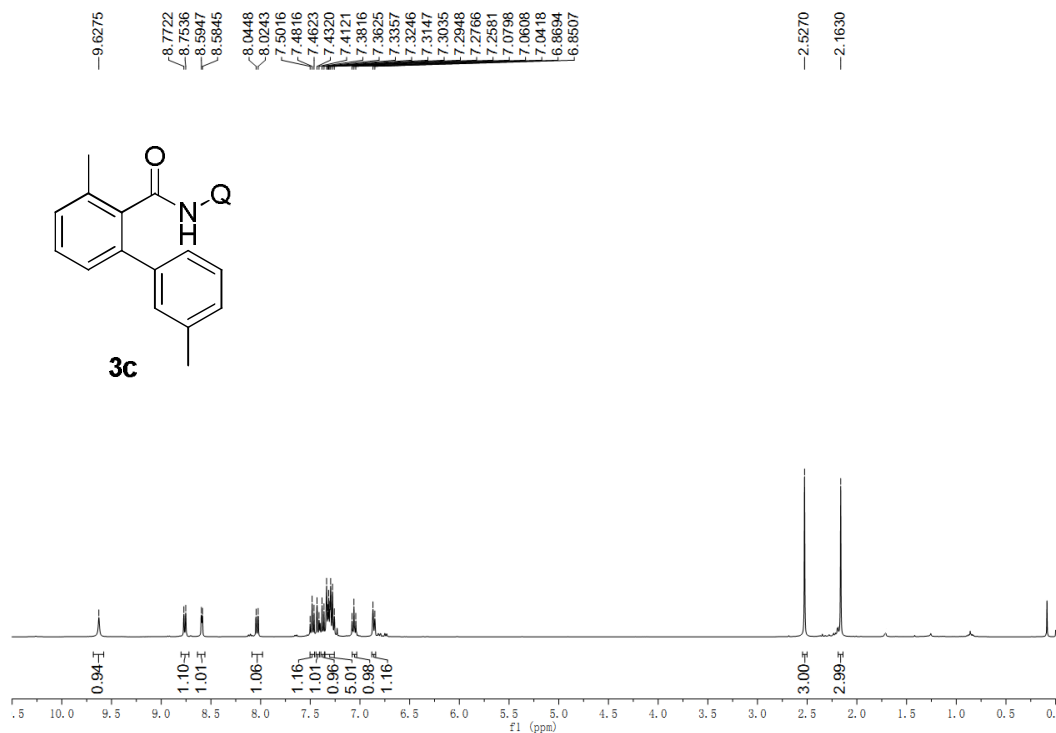
1. A. M. Suess, M. Z. Ertem, C. J. Cramer and S. S. Stahl, *J. Am. Chem. Soc.*, 2013, **135**, 9797.
2. (a) L. D. Tran, I. Popov and O. Daugulis, *J. Am. Chem. Soc.*, 2012, **134**, 18237; (b) T. Truong, K. Klimovica and O. Daugulis, *J. Am. Chem. Soc.*, 2013, **135**, 9342; (c) L. D. Tran, J. Roane and O. Daugulis, *Angew. Chem., Int. Ed.*, 2013, **52**, 6043; (d) J. Roane and O. Daugulis, *Org. Lett.*,

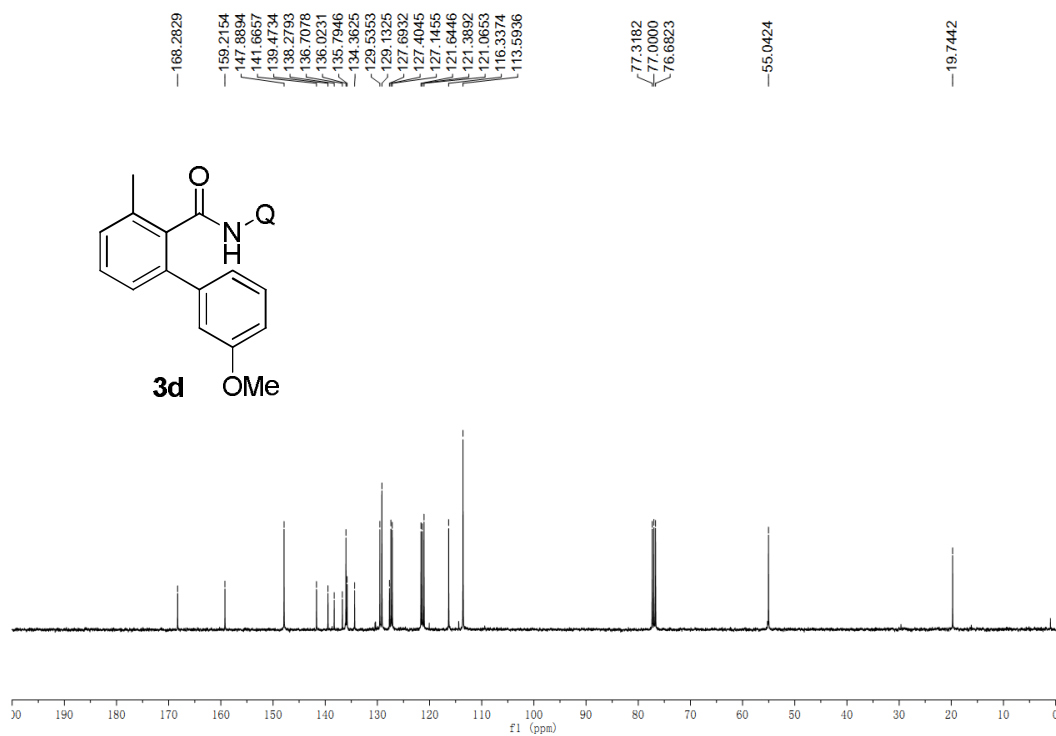
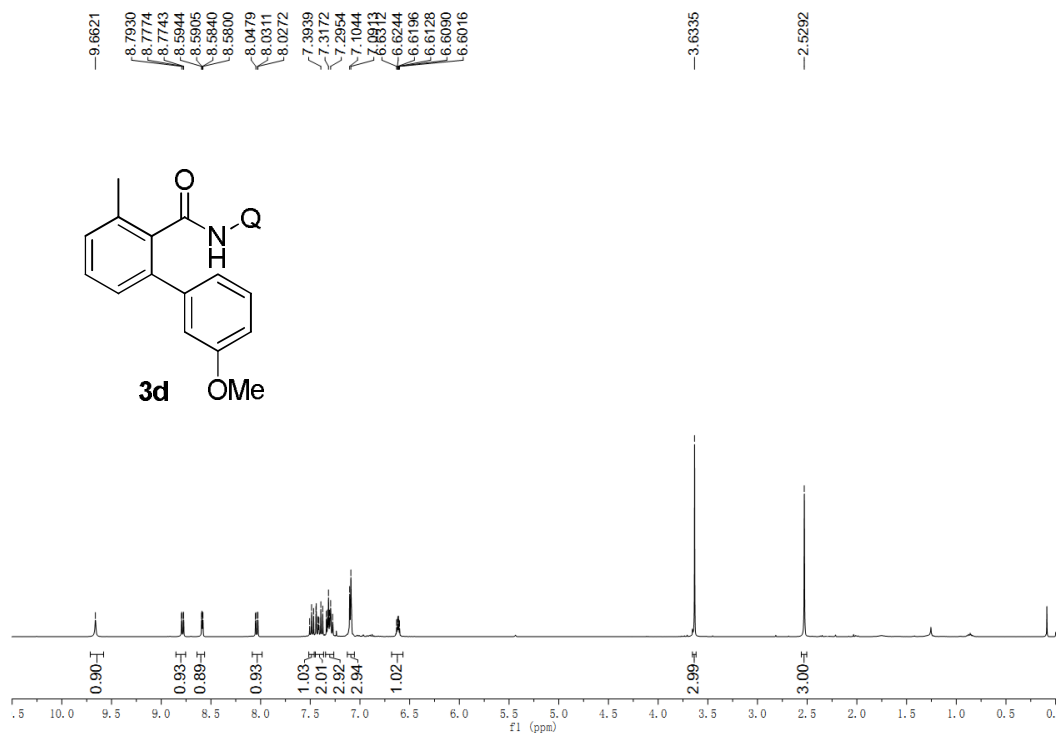
2013, **15**, 5842; (e) M. Nishino, K. Hirano, T. Satoh and M. Miura, *Angew. Chem., Int. Ed.*, 2013, **52**, 4457.

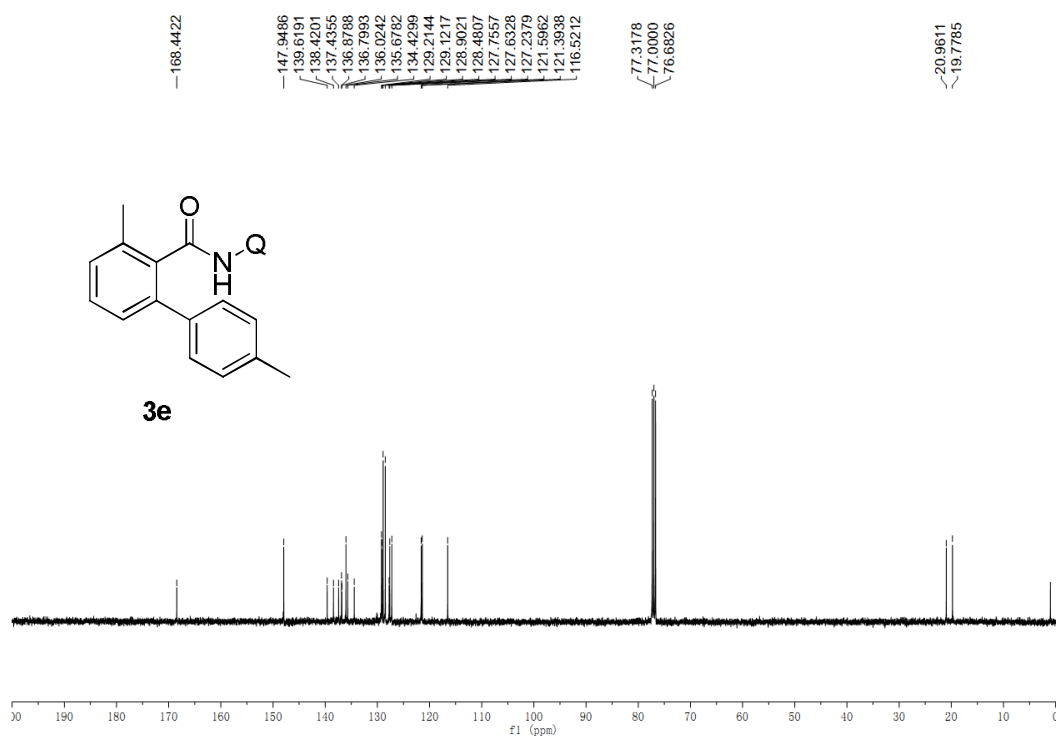
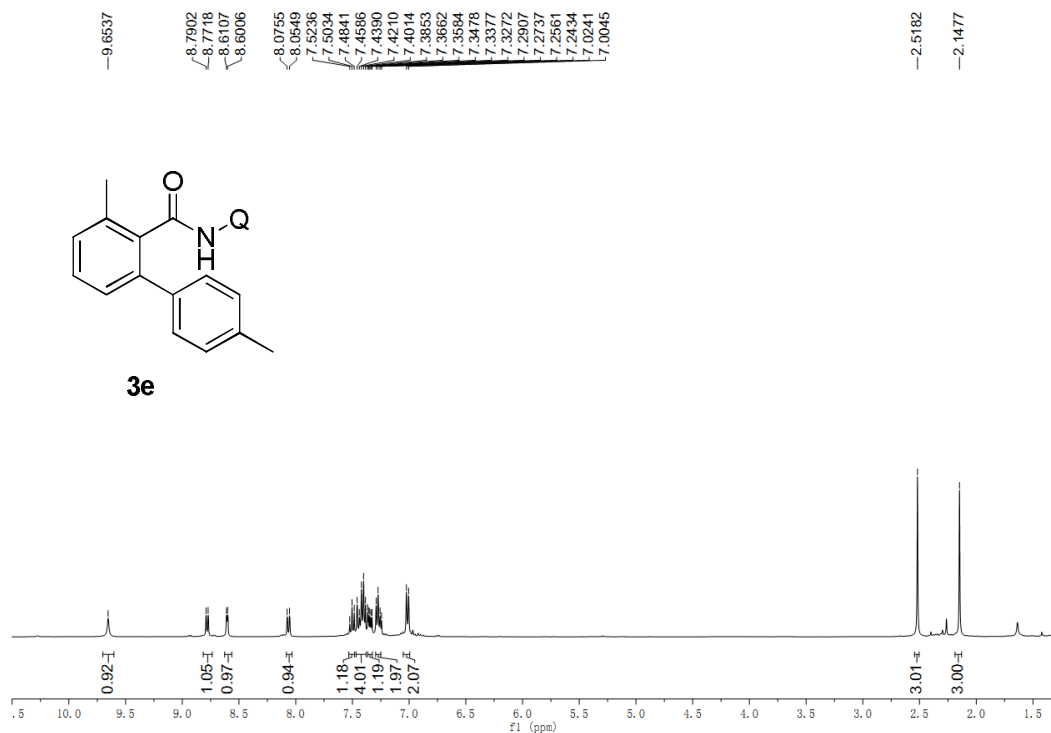
3. A. Yokota, Y. Aihara and N. Chatani, *J. Org. Chem.*, 2014, **79**, 11922.

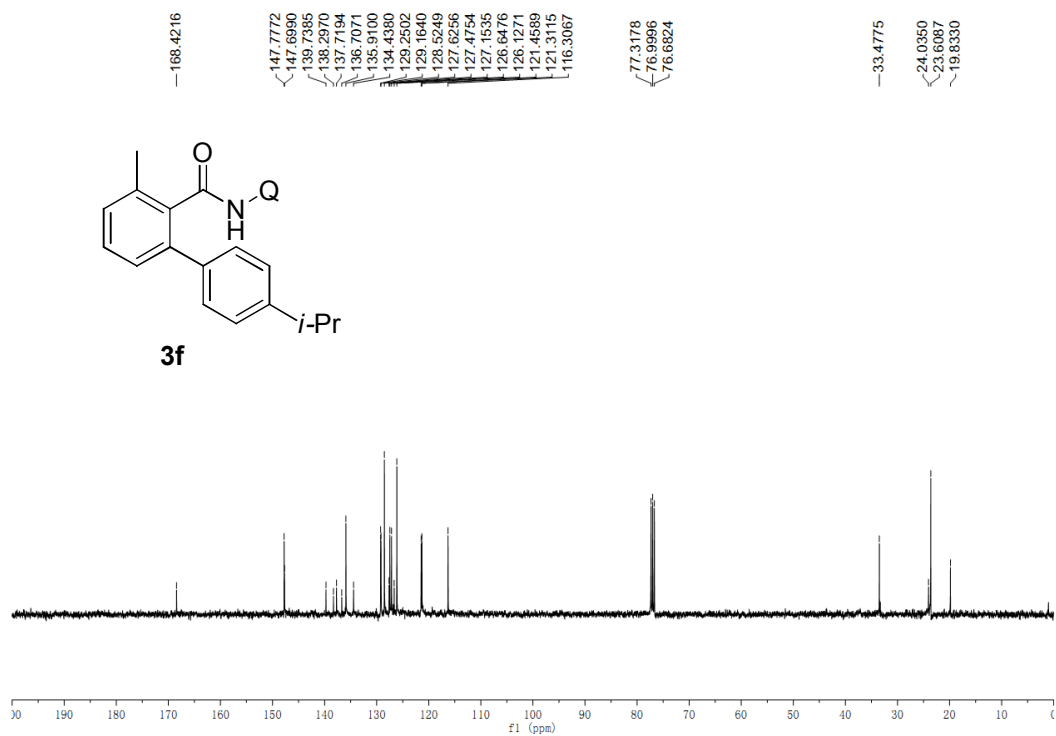
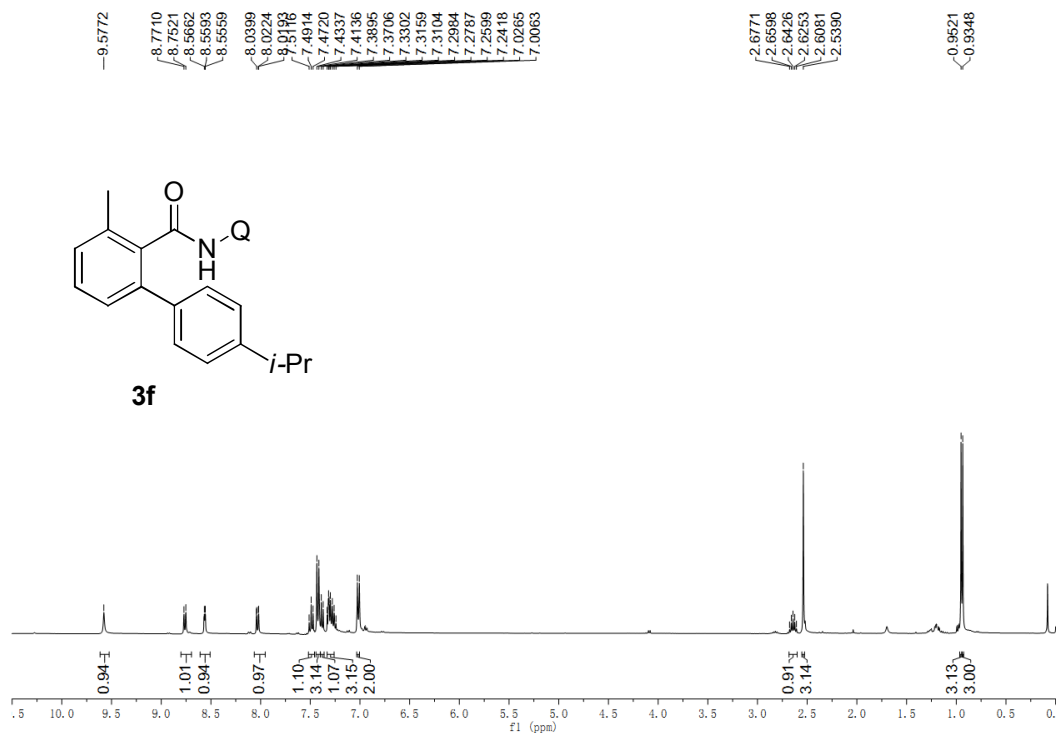


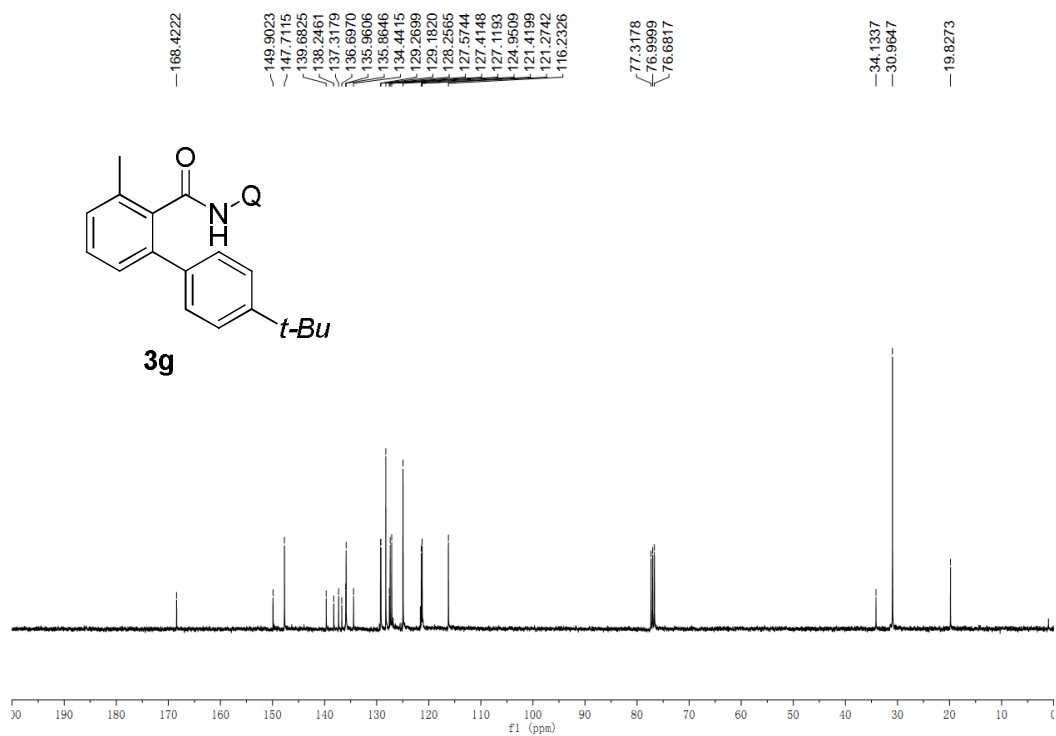
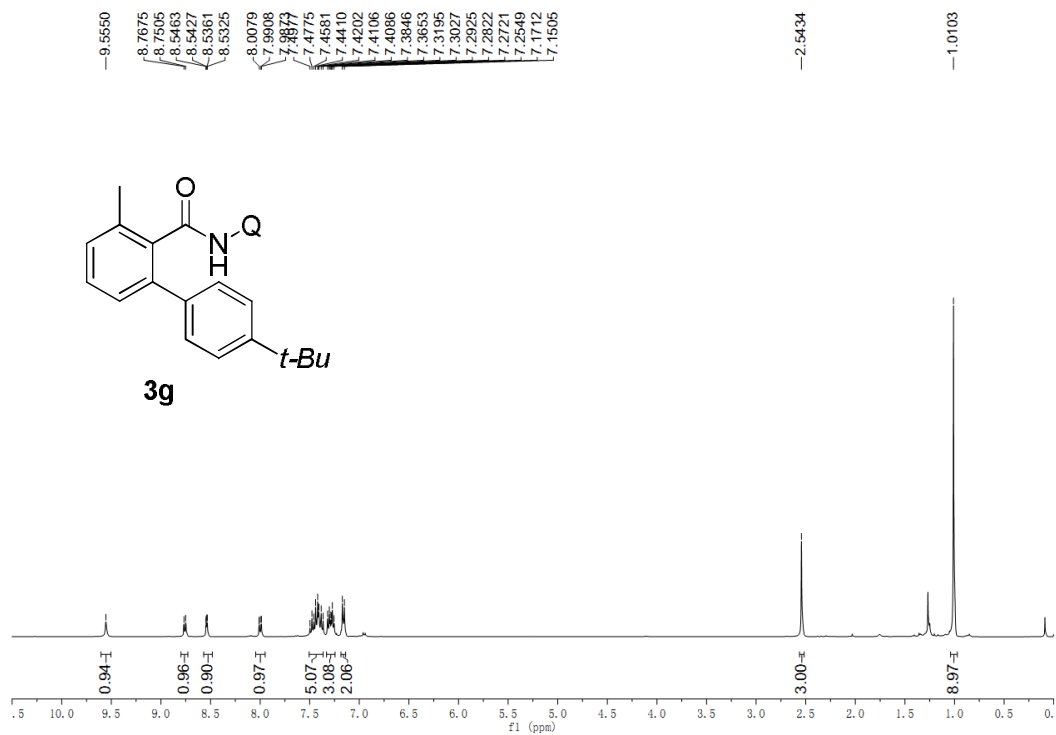


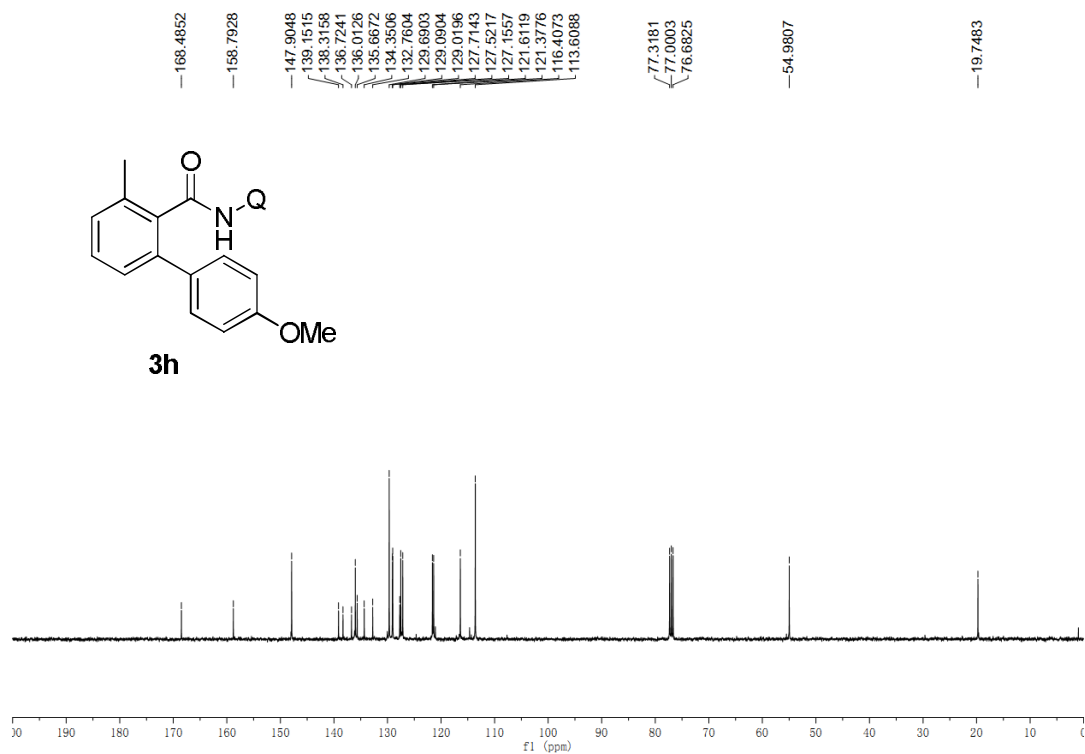
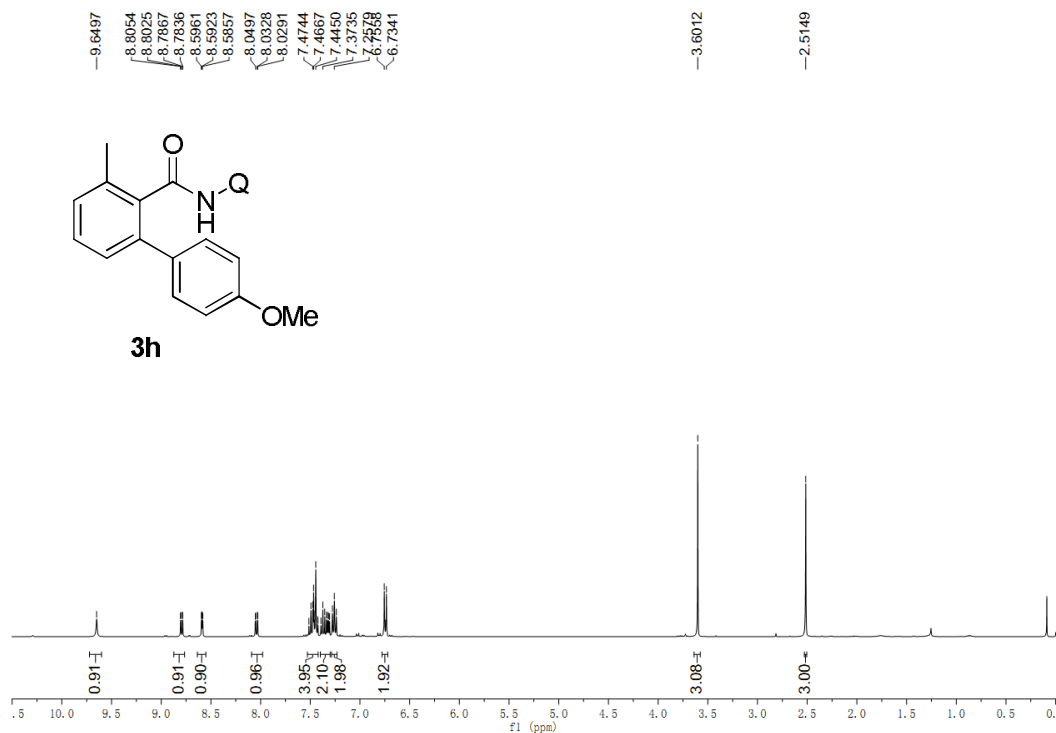


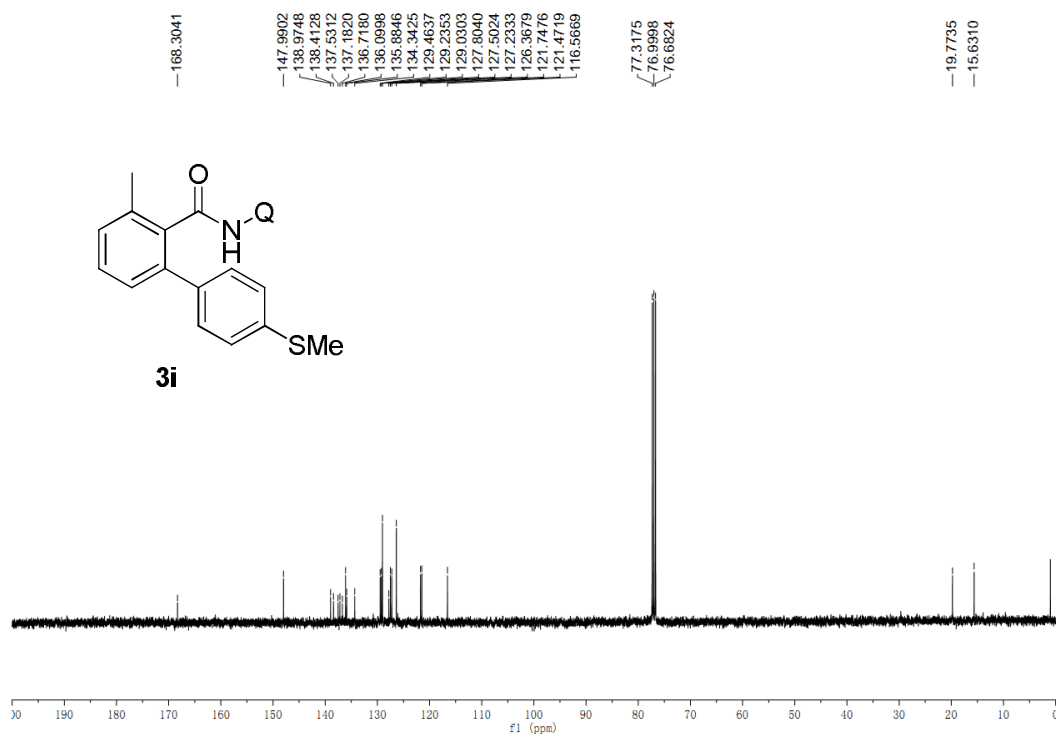
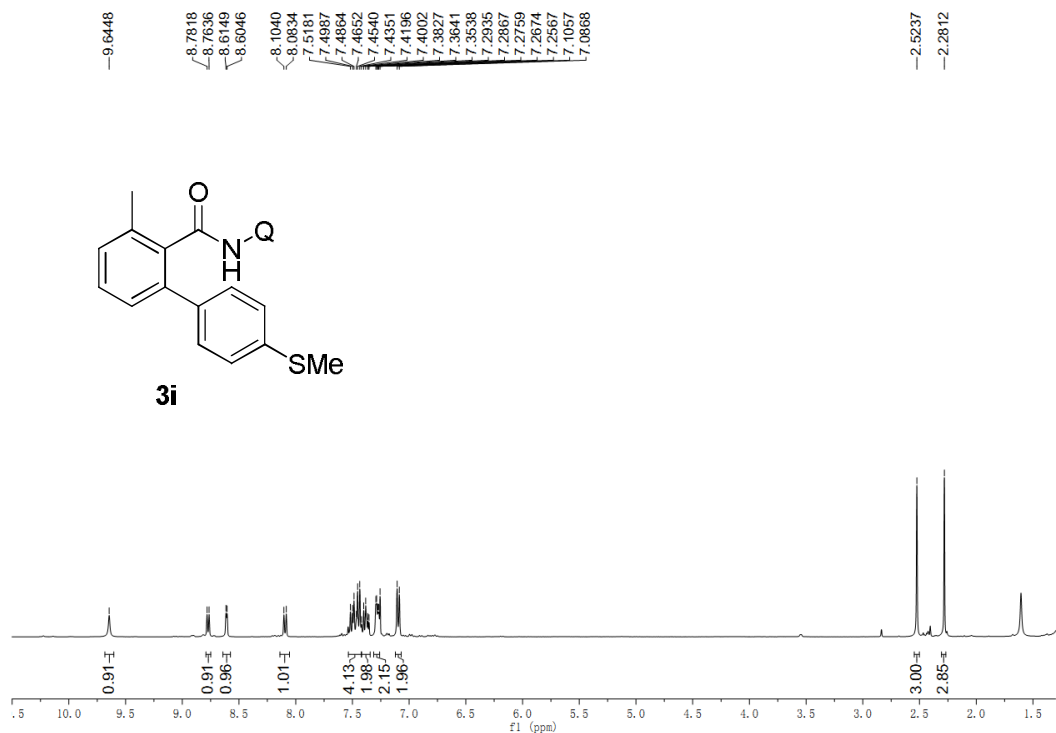






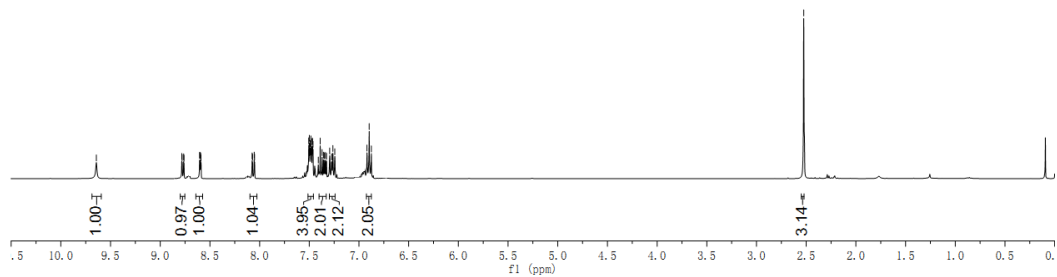
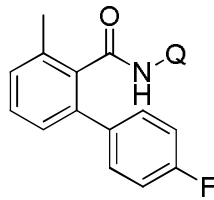






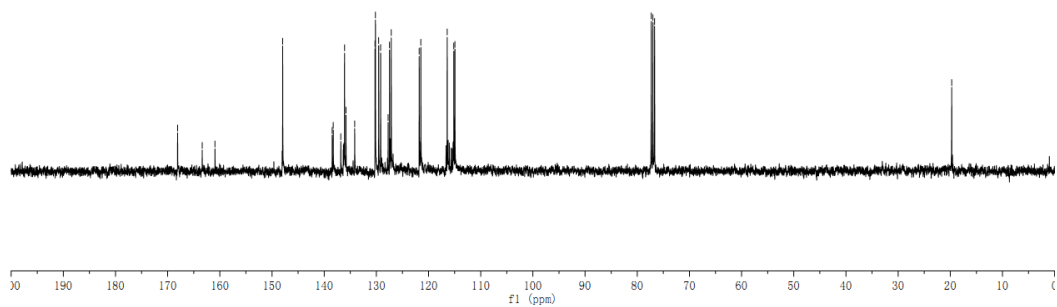
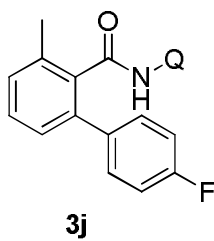
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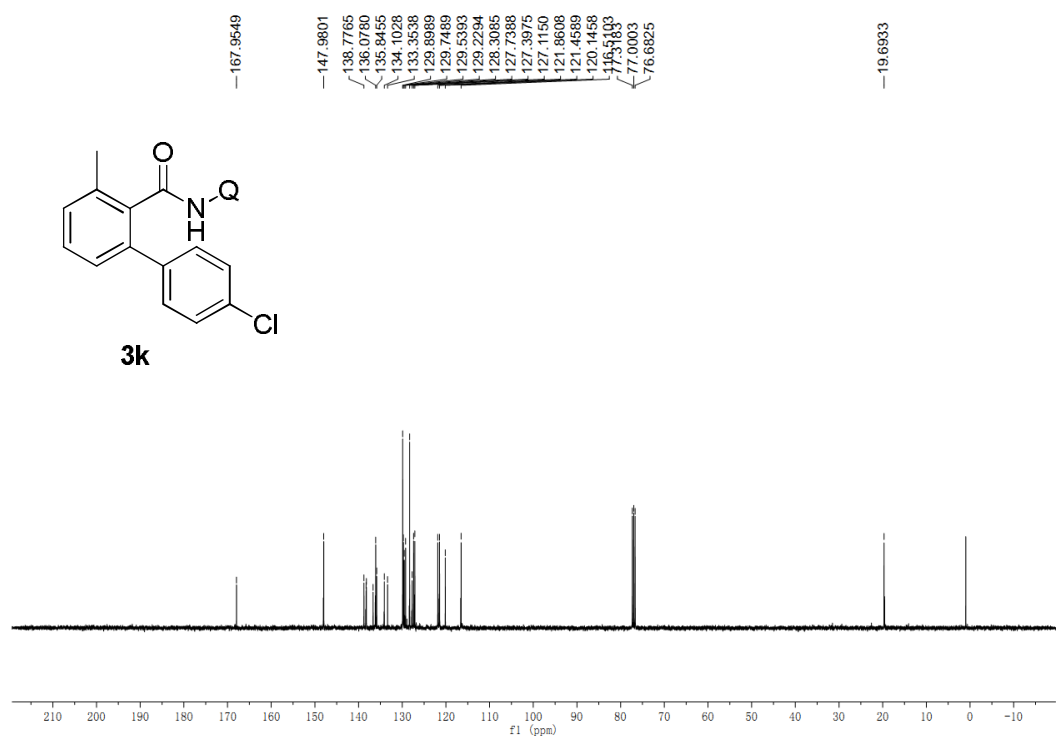
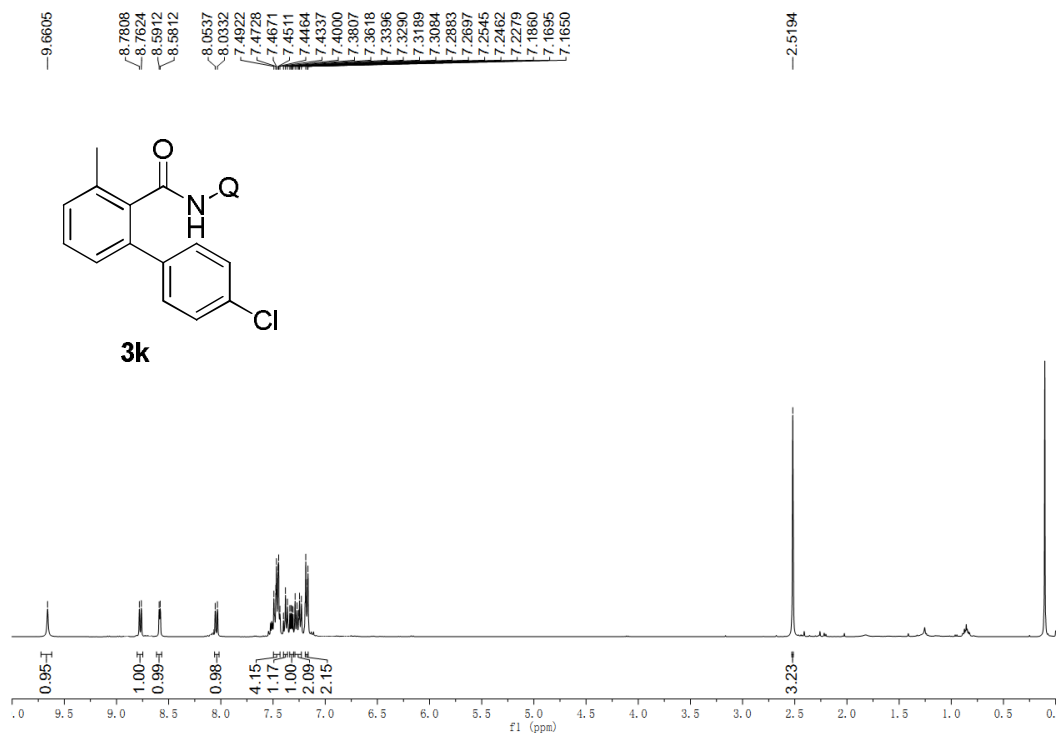
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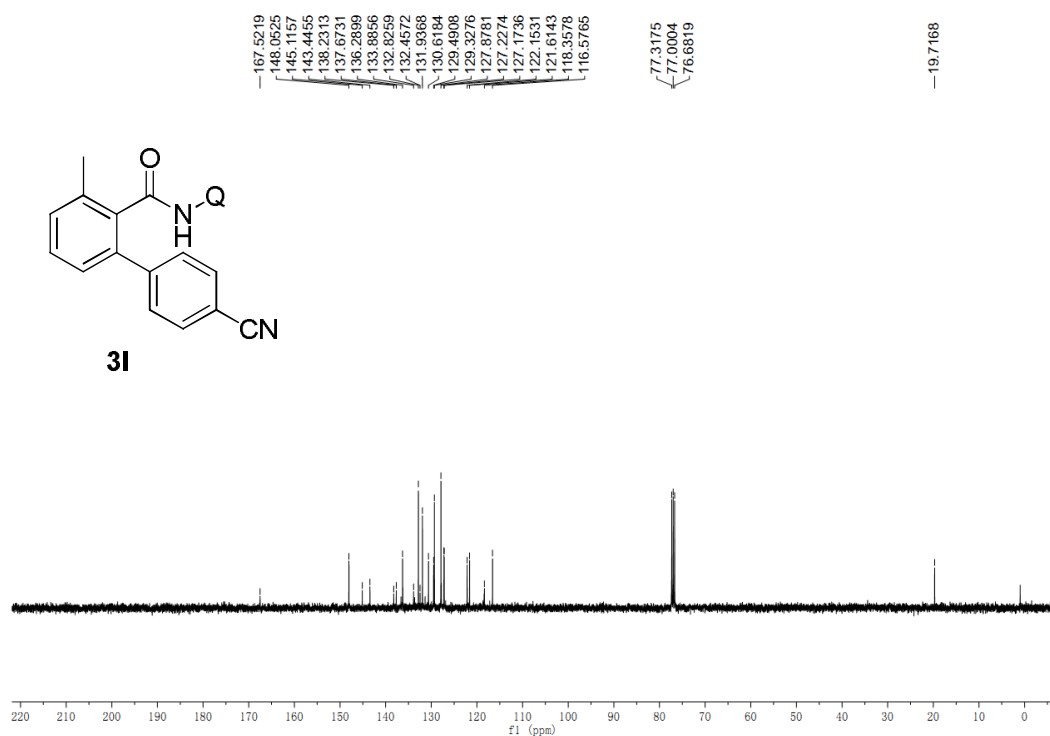
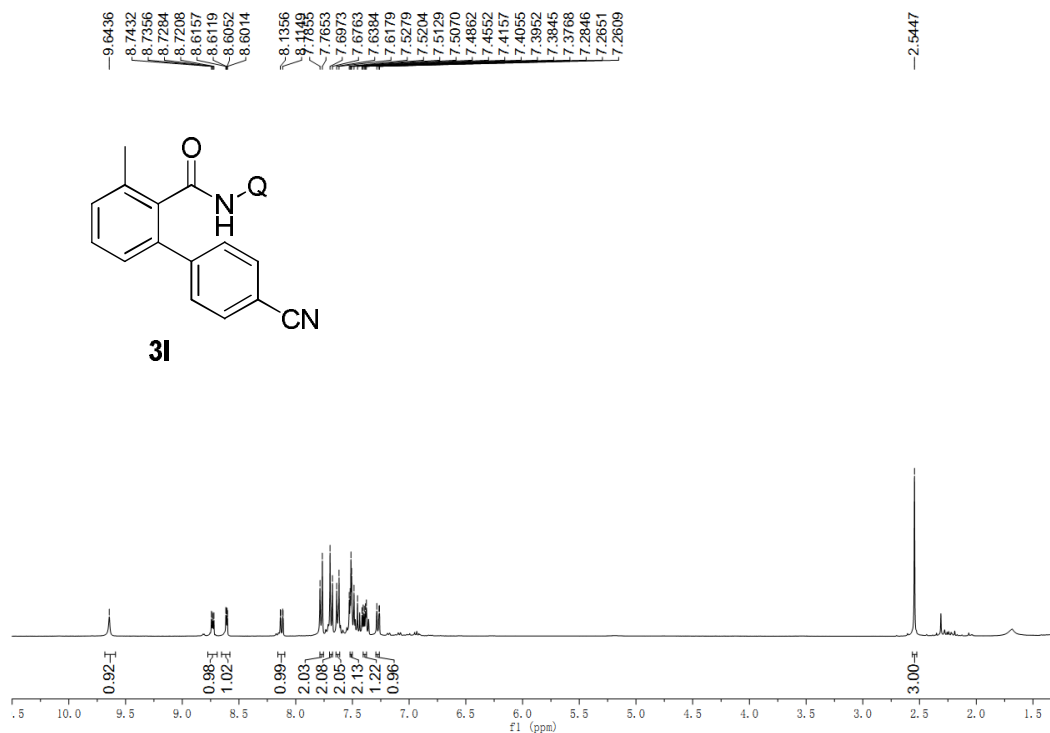


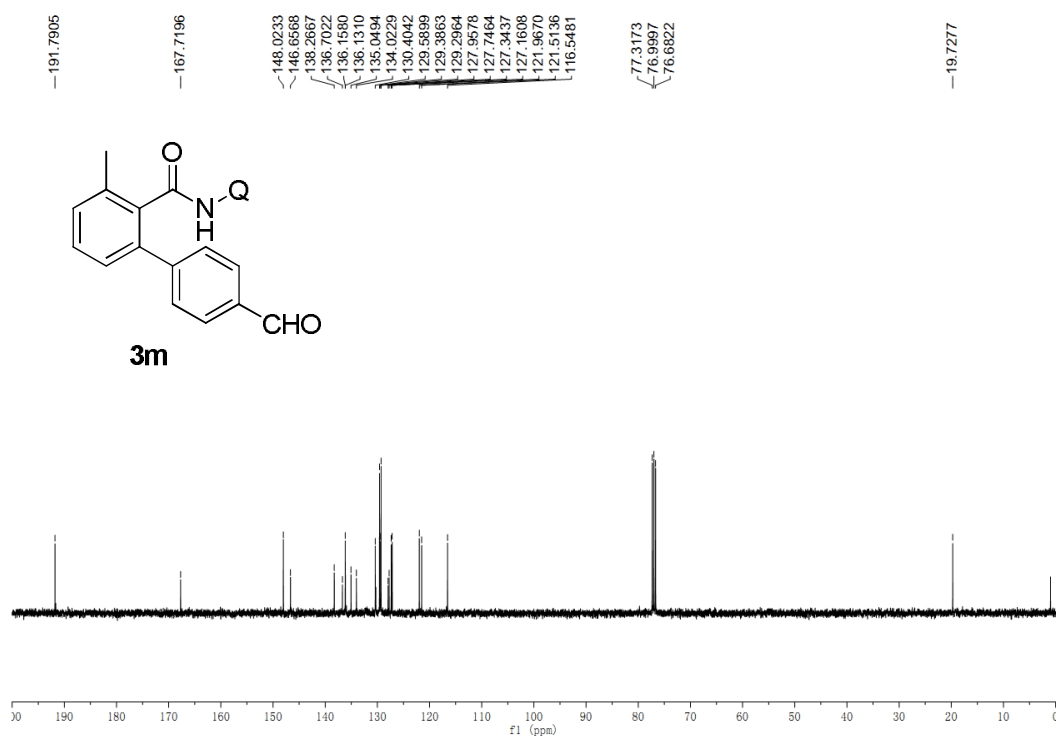
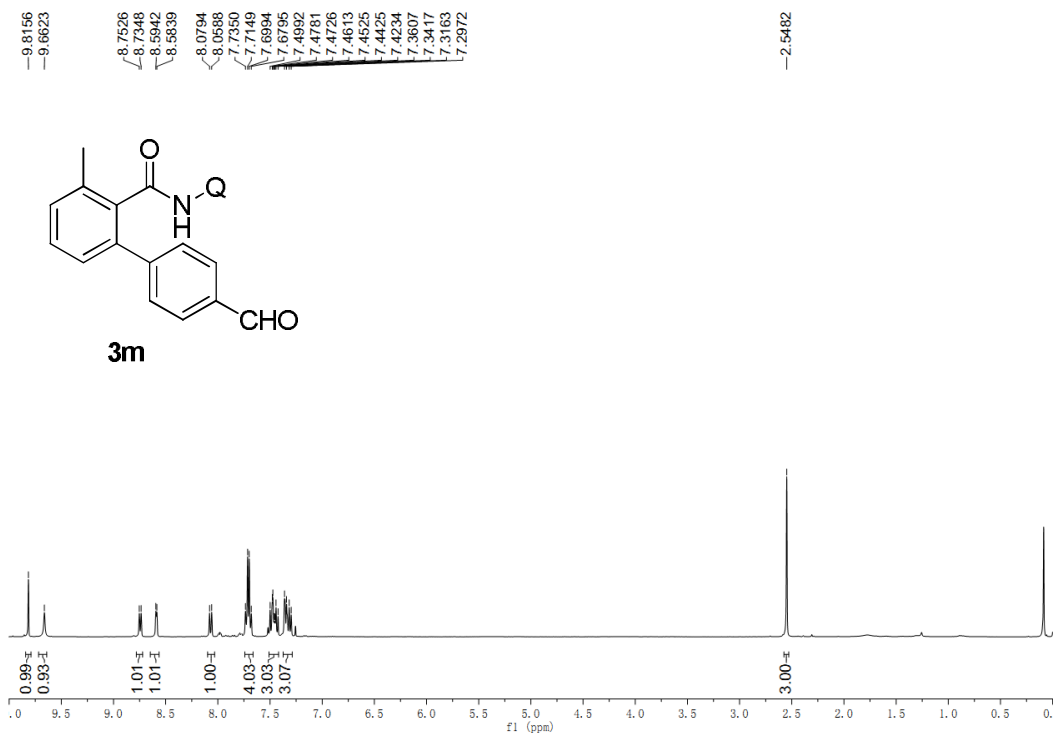
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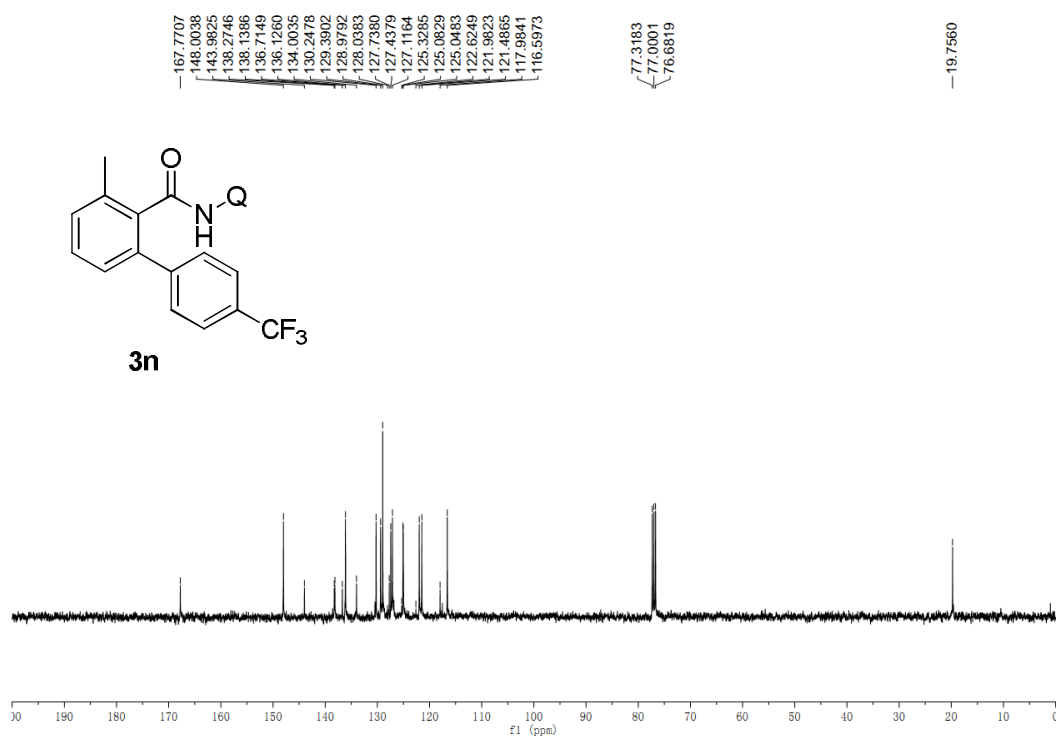
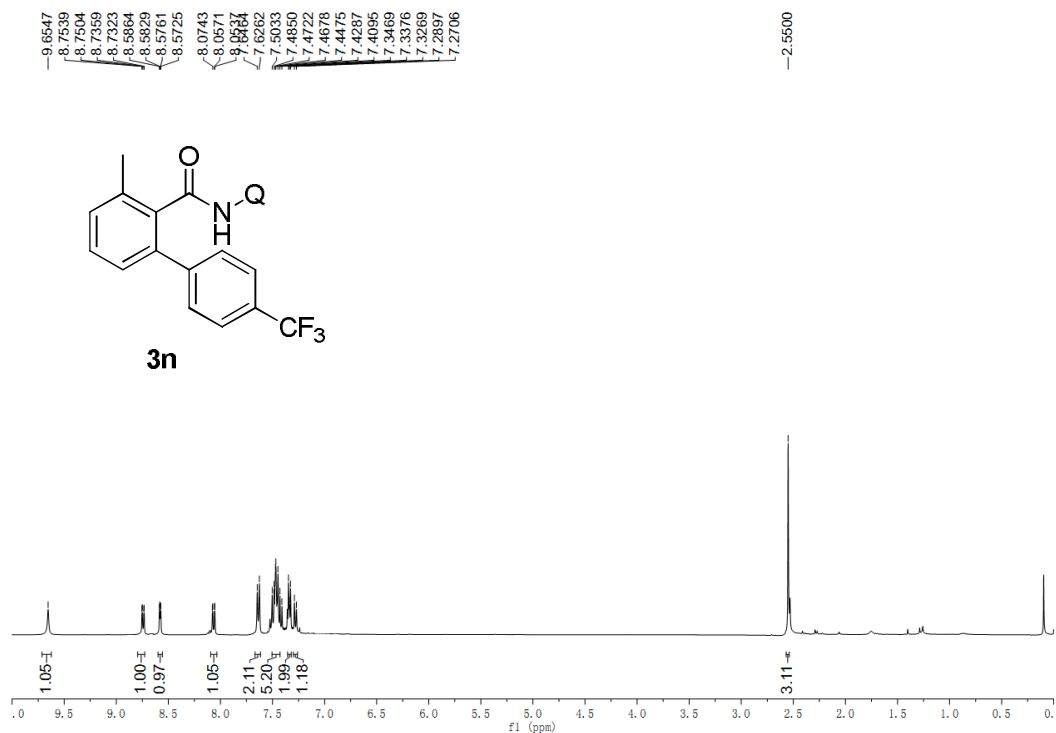
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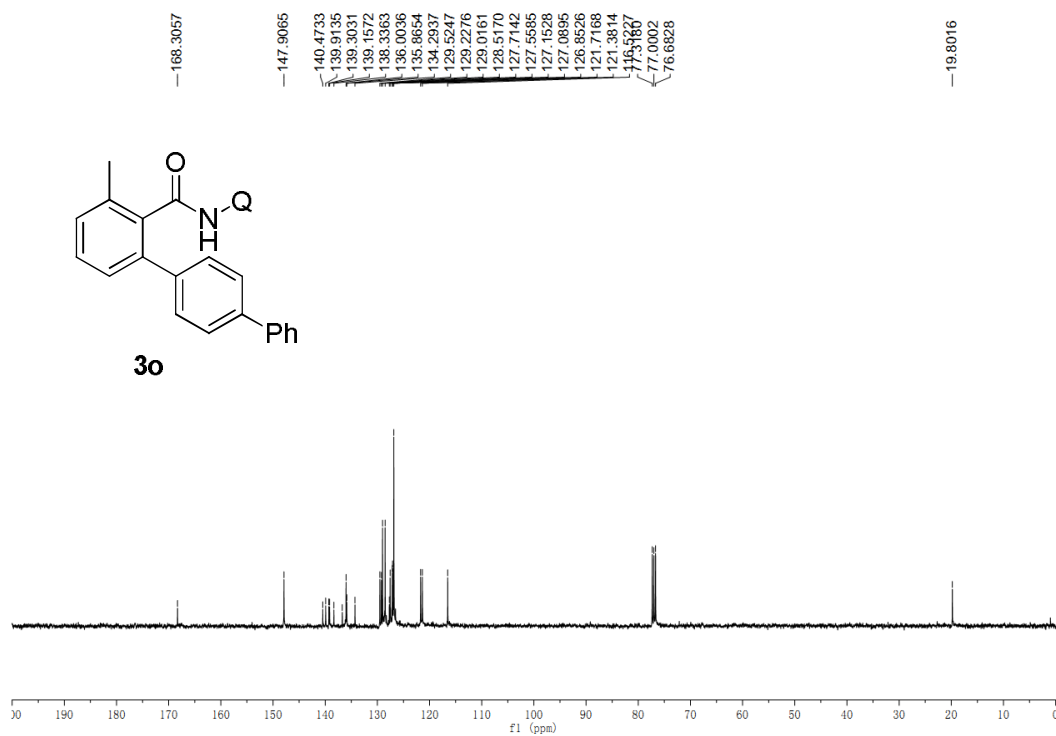
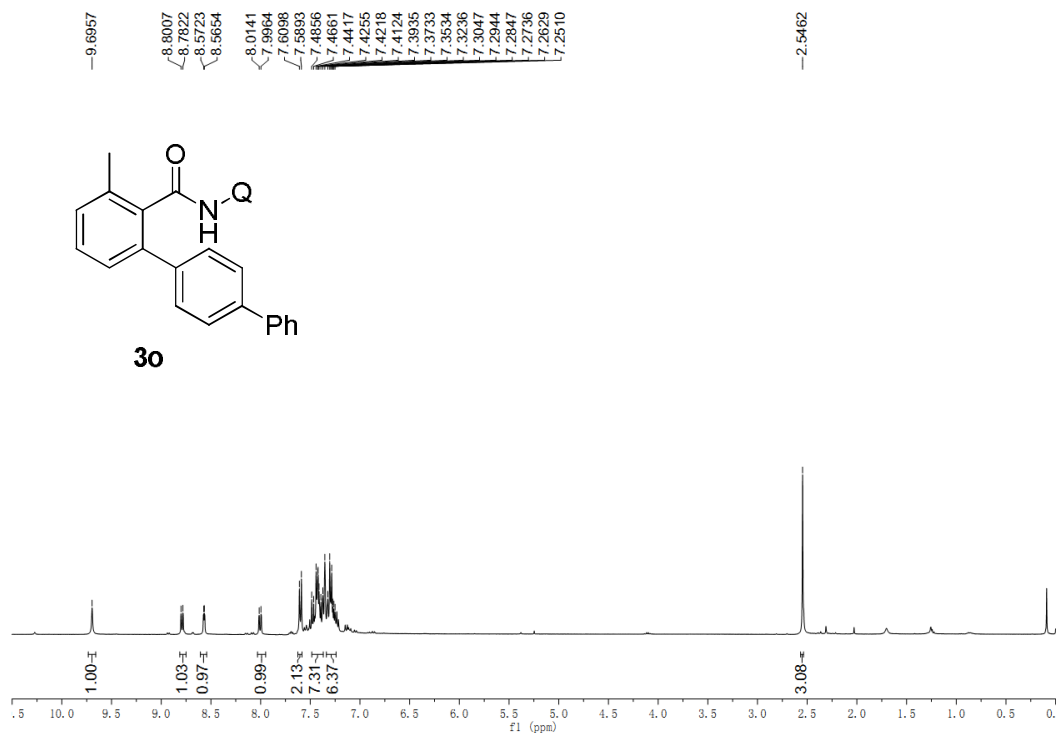


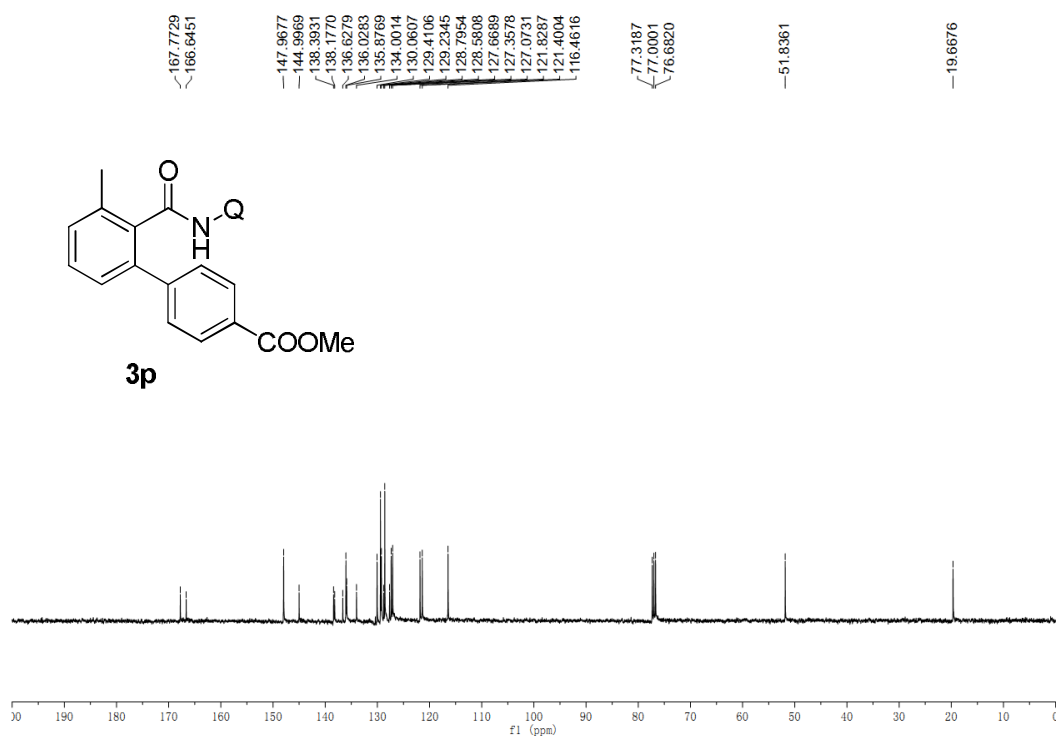
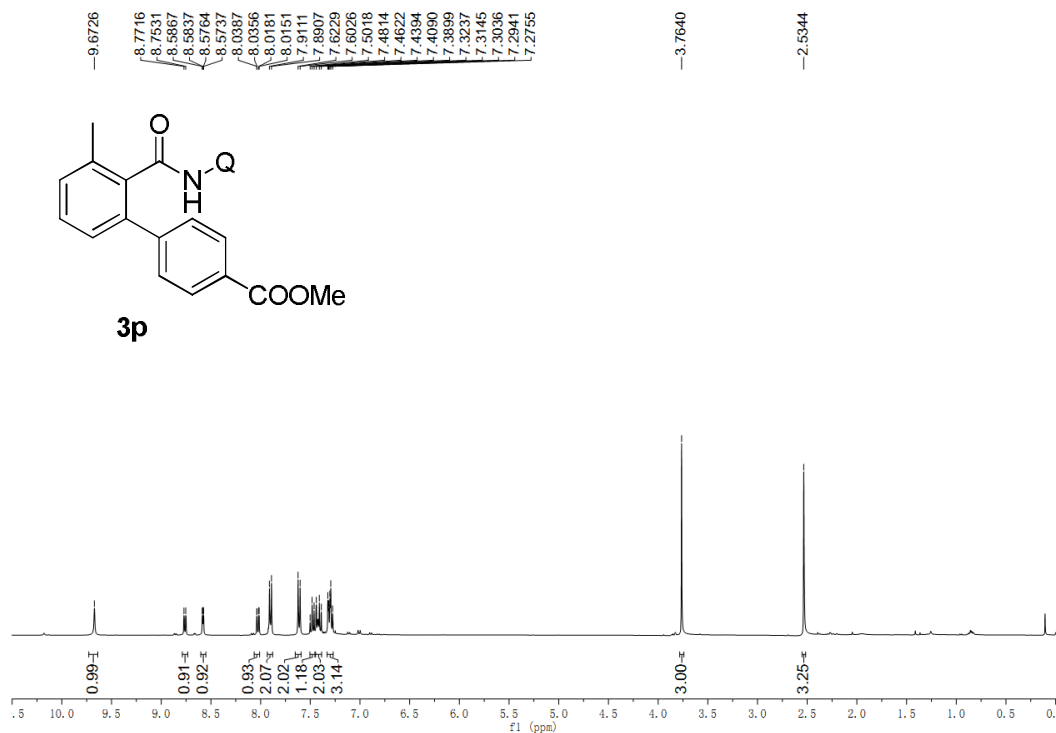


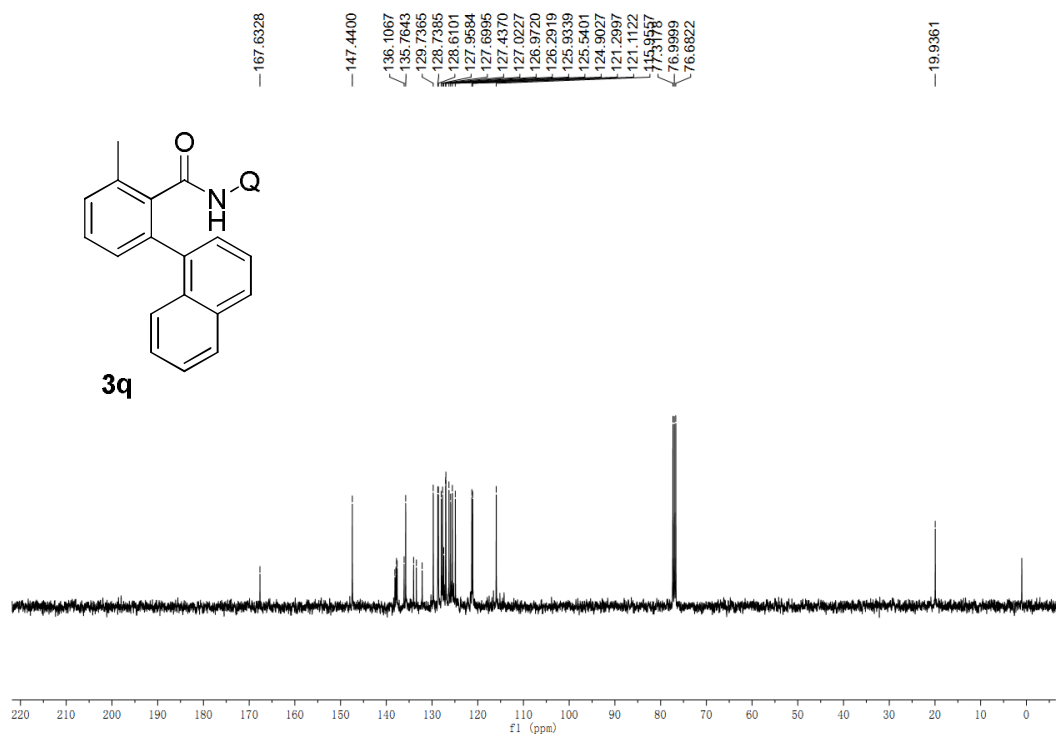
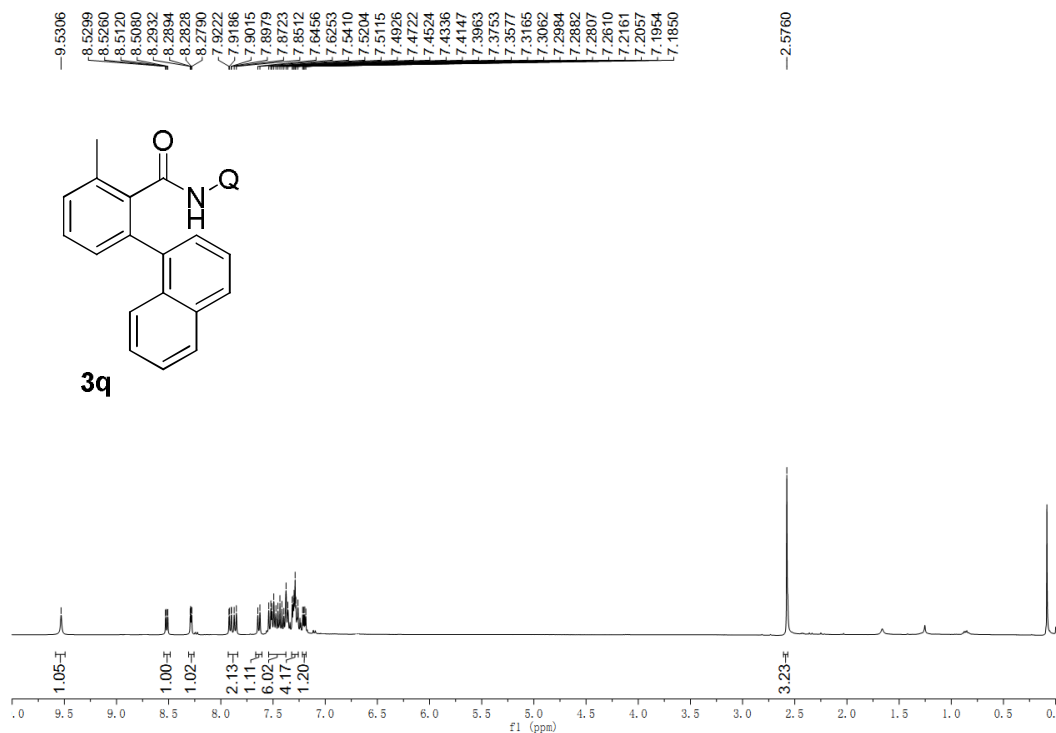


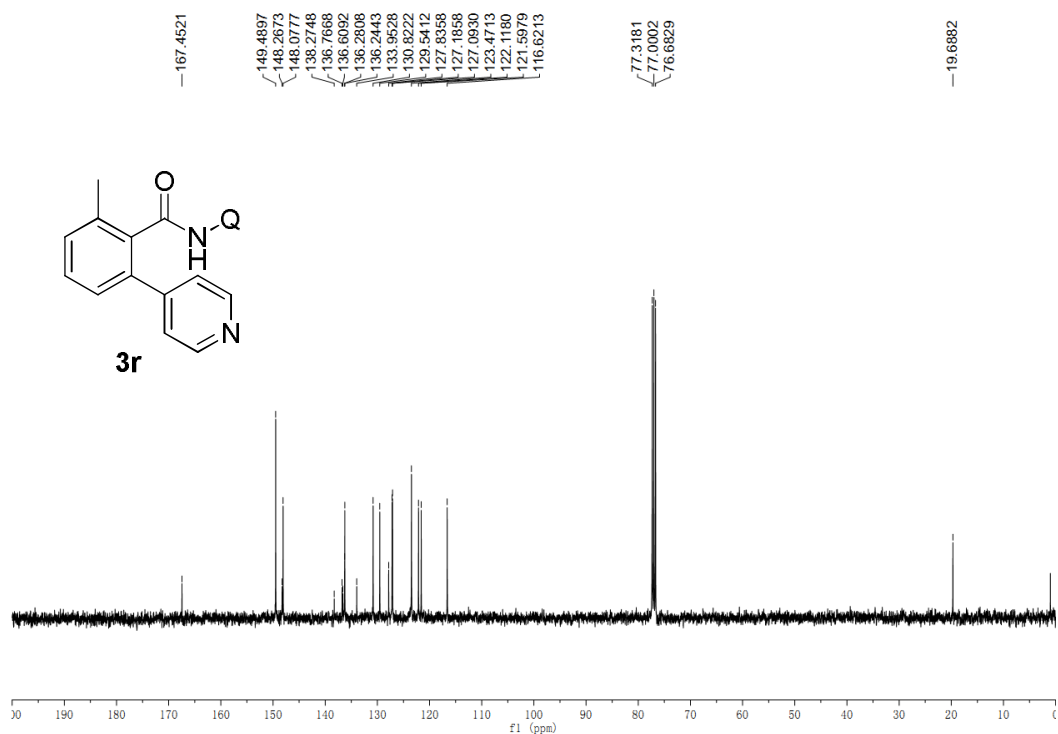
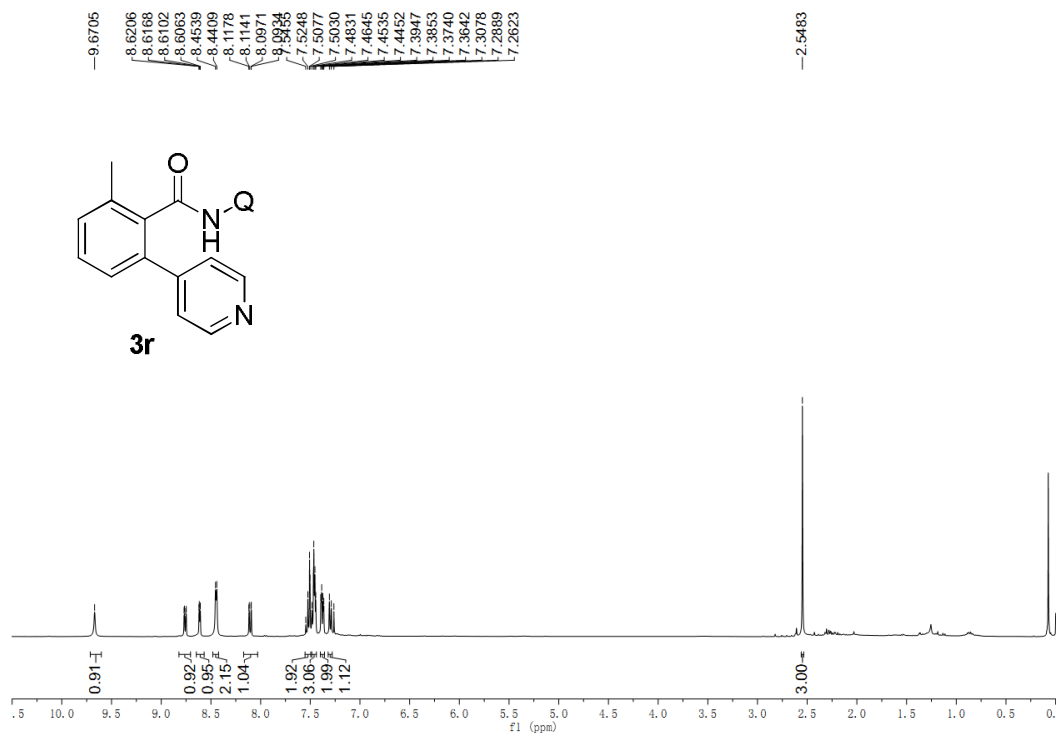


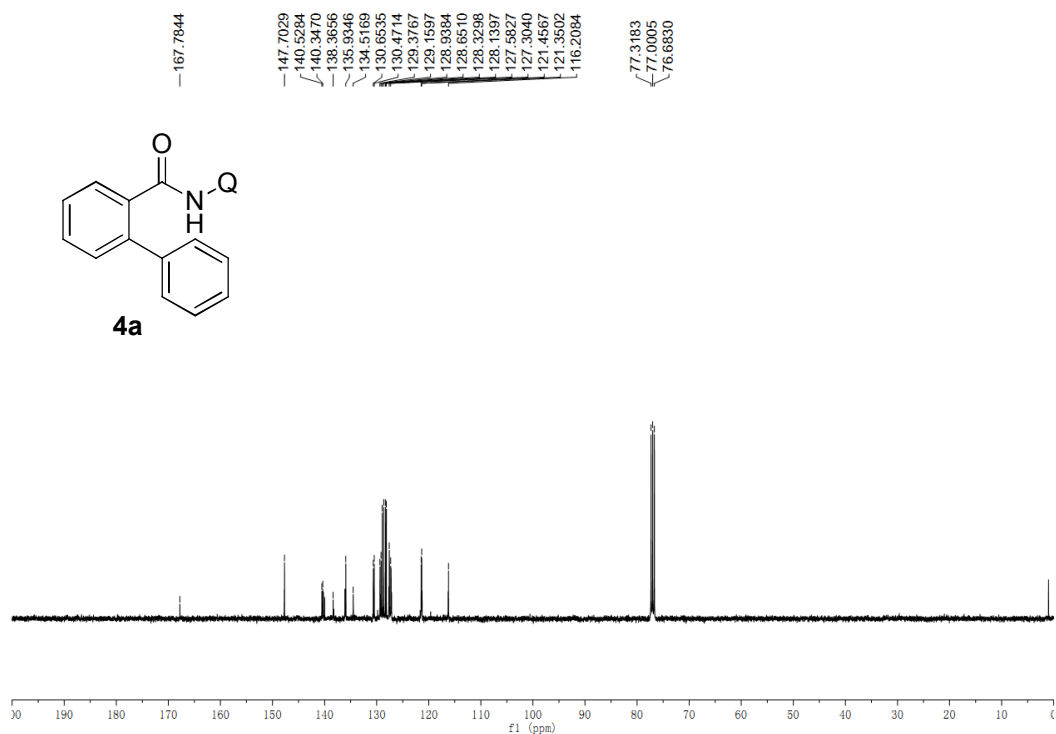
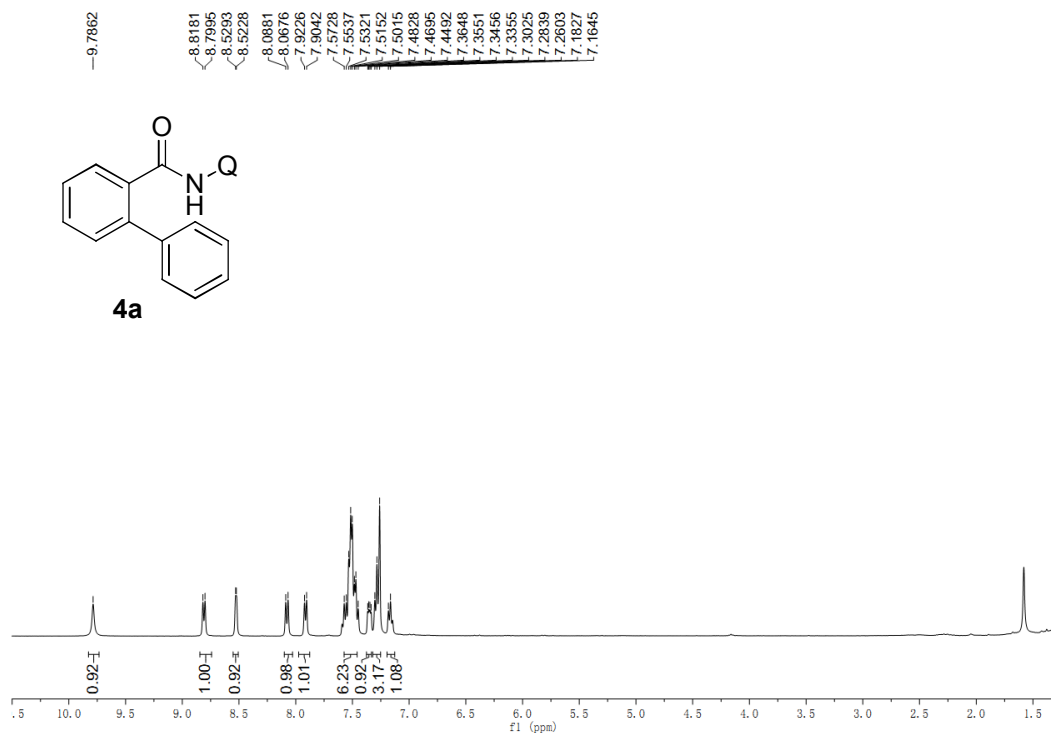


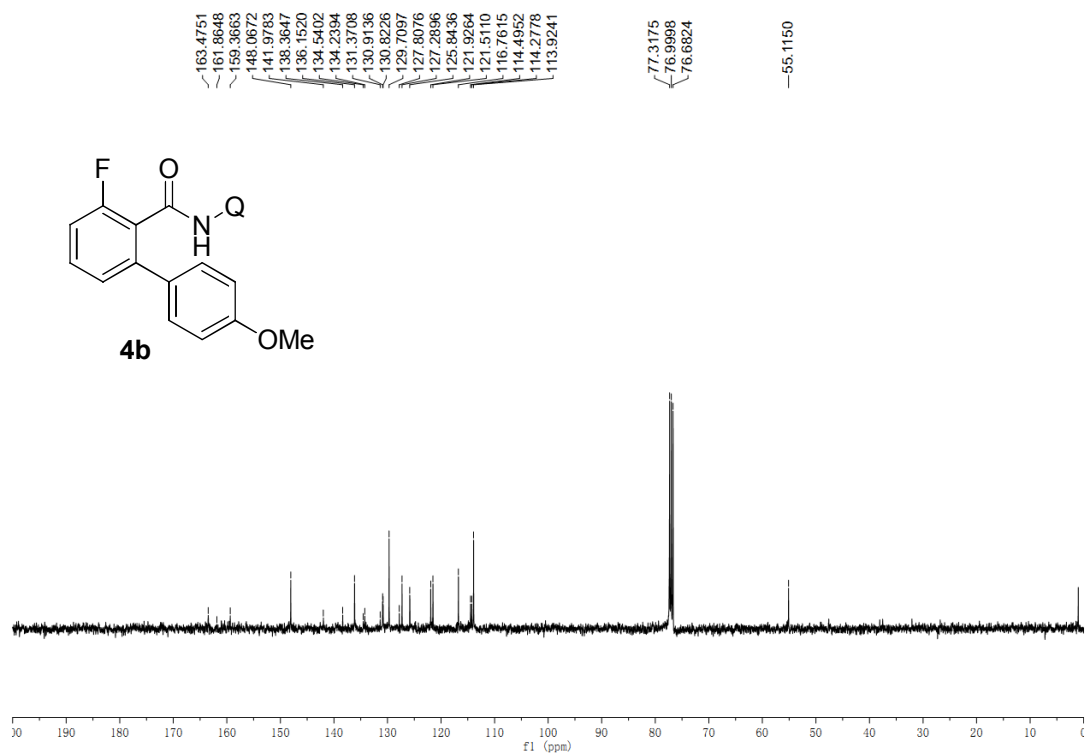
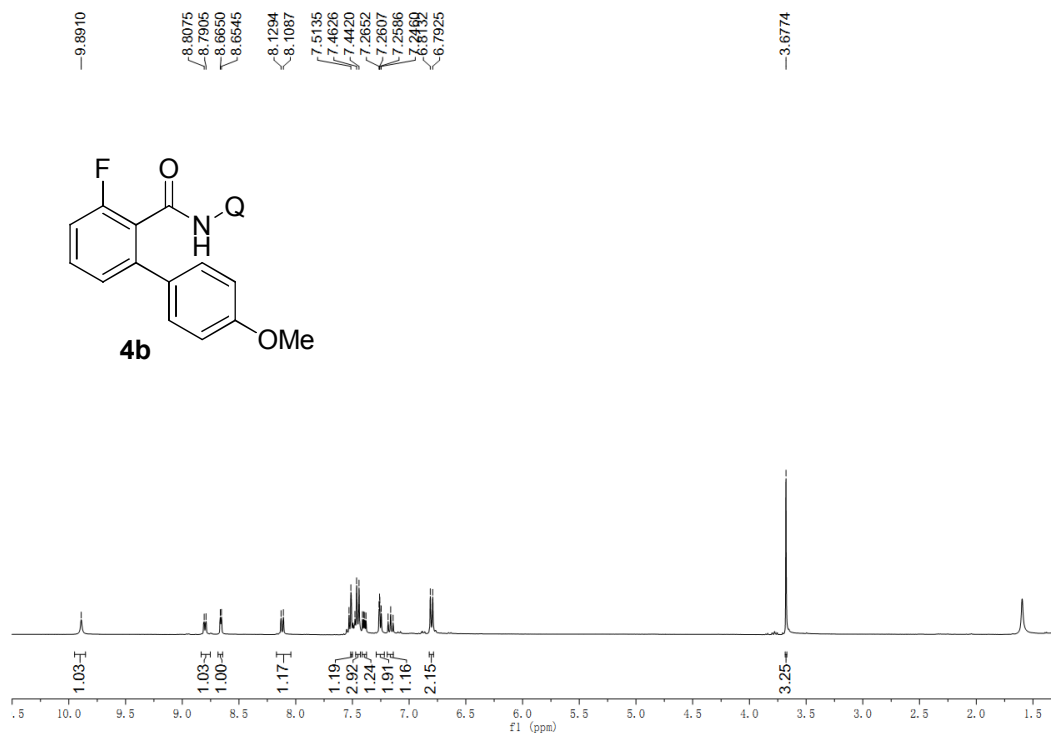


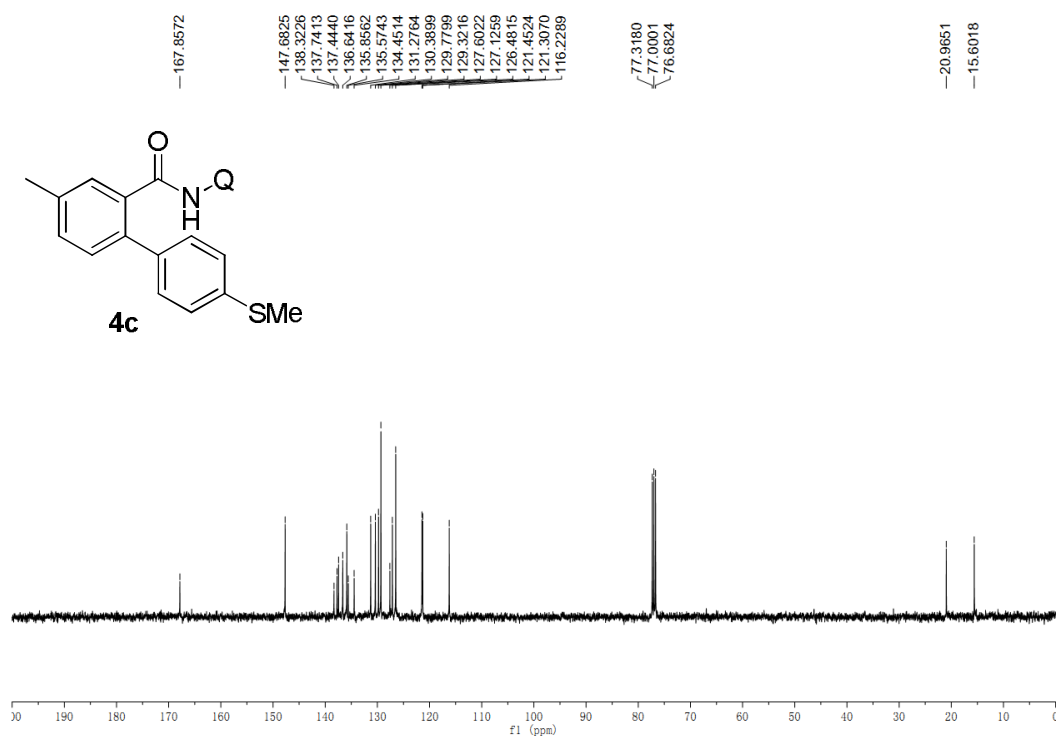
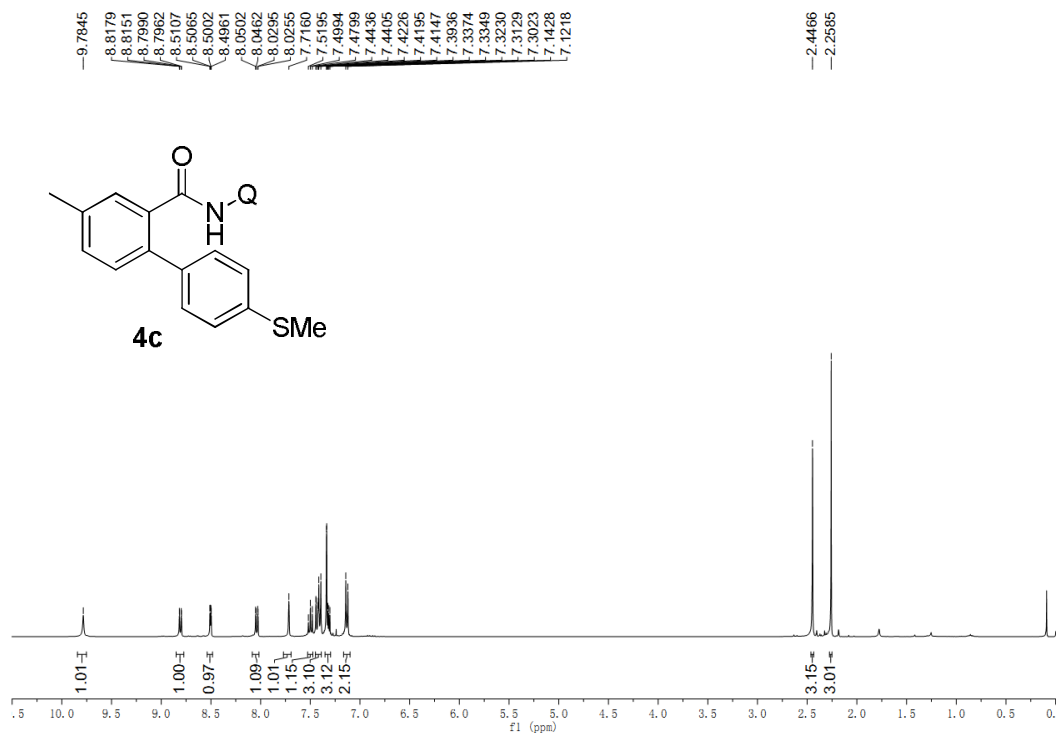


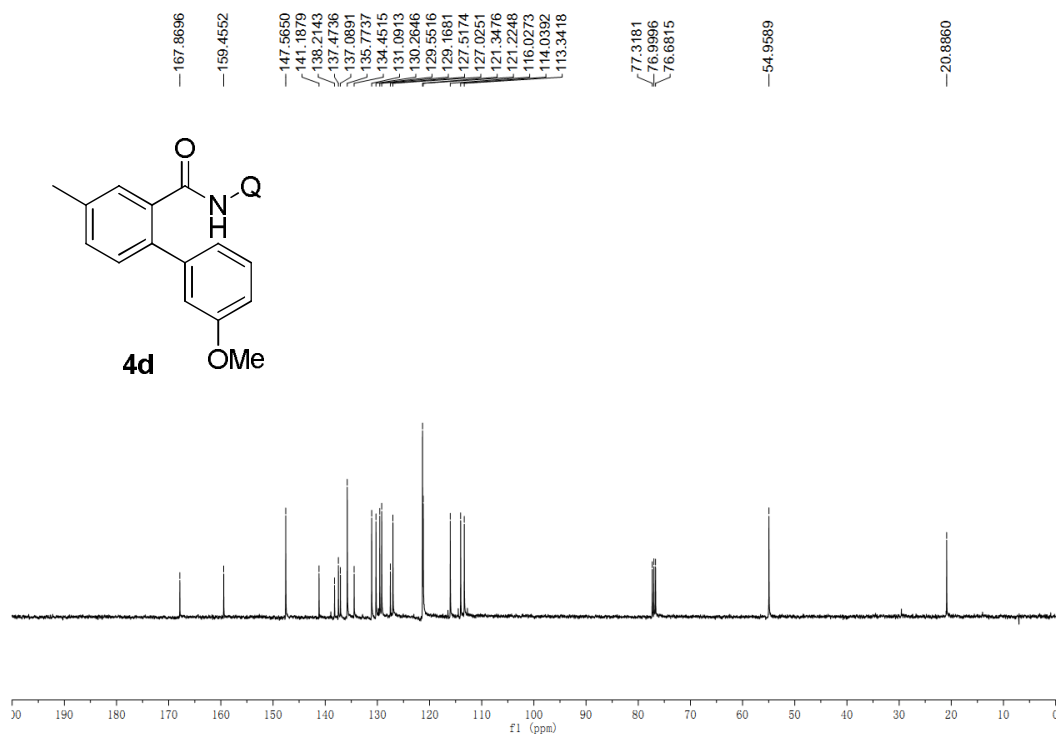
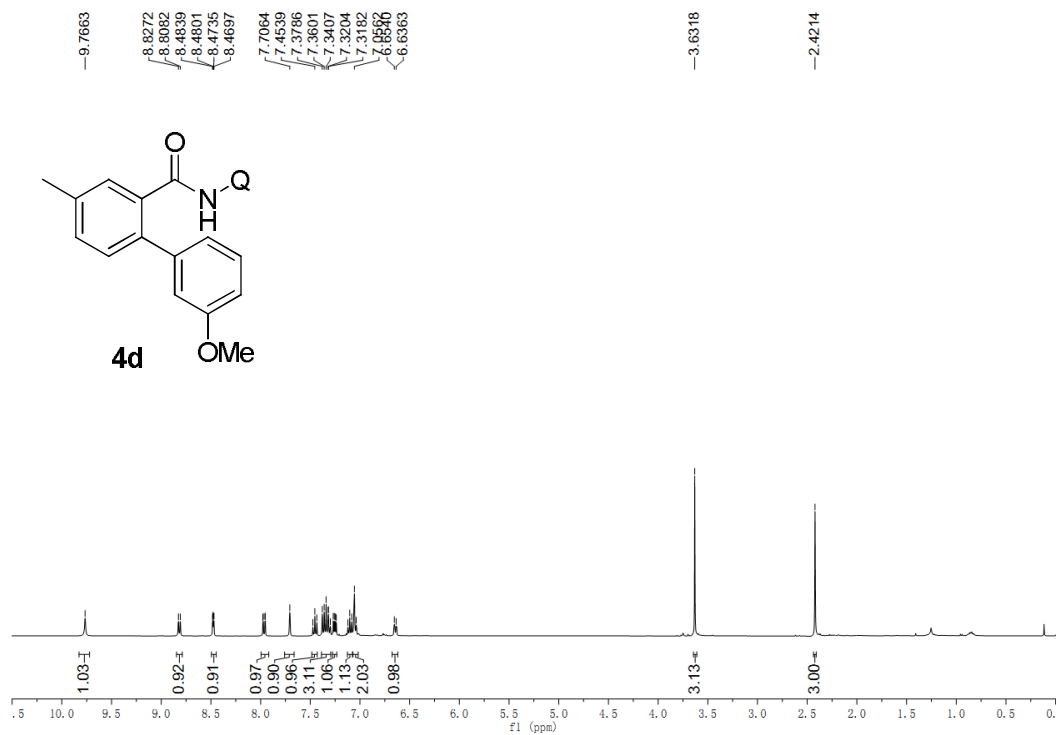


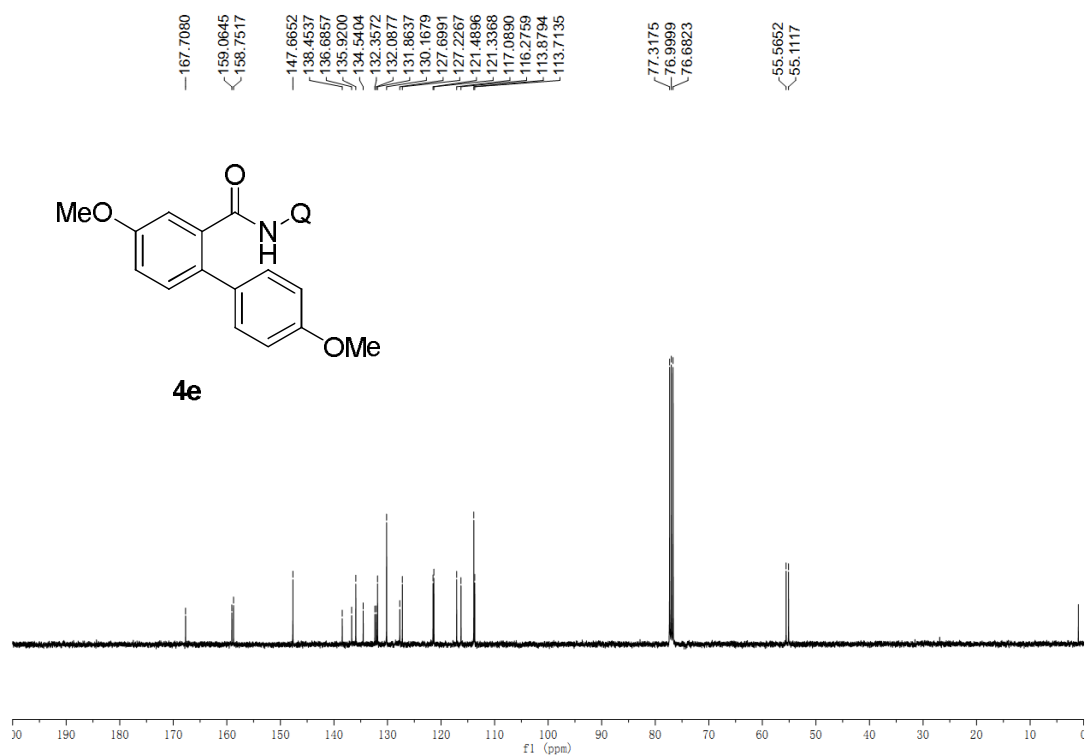
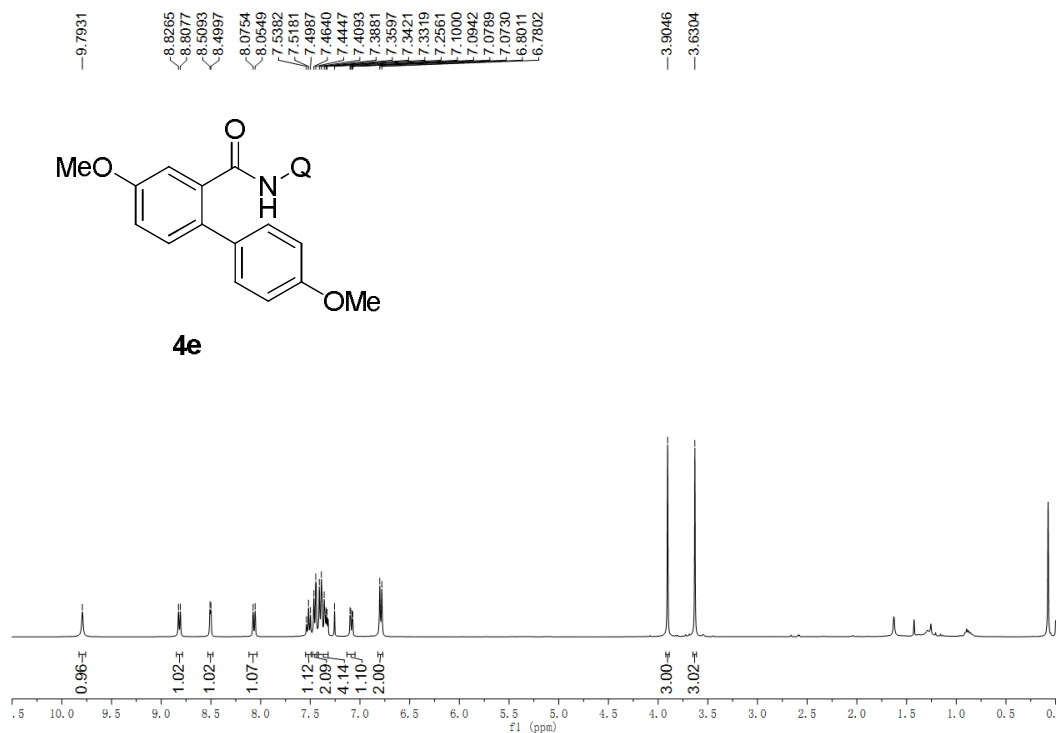




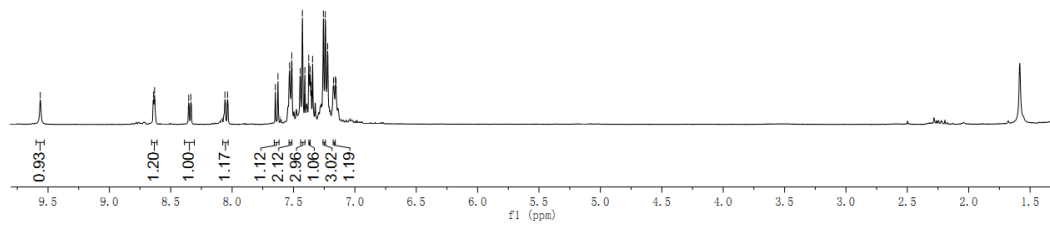
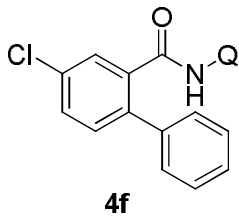




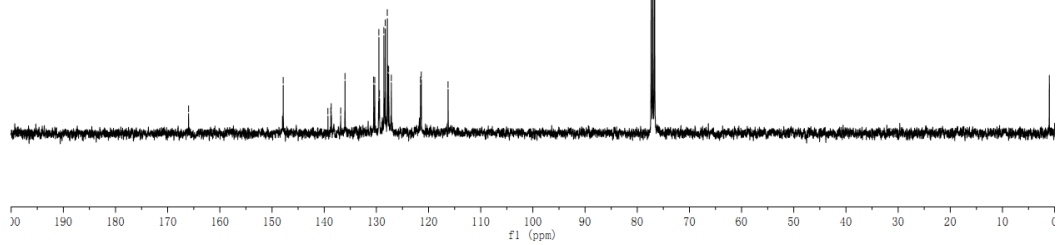
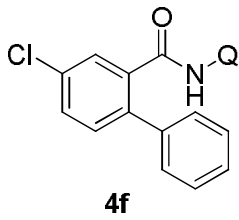


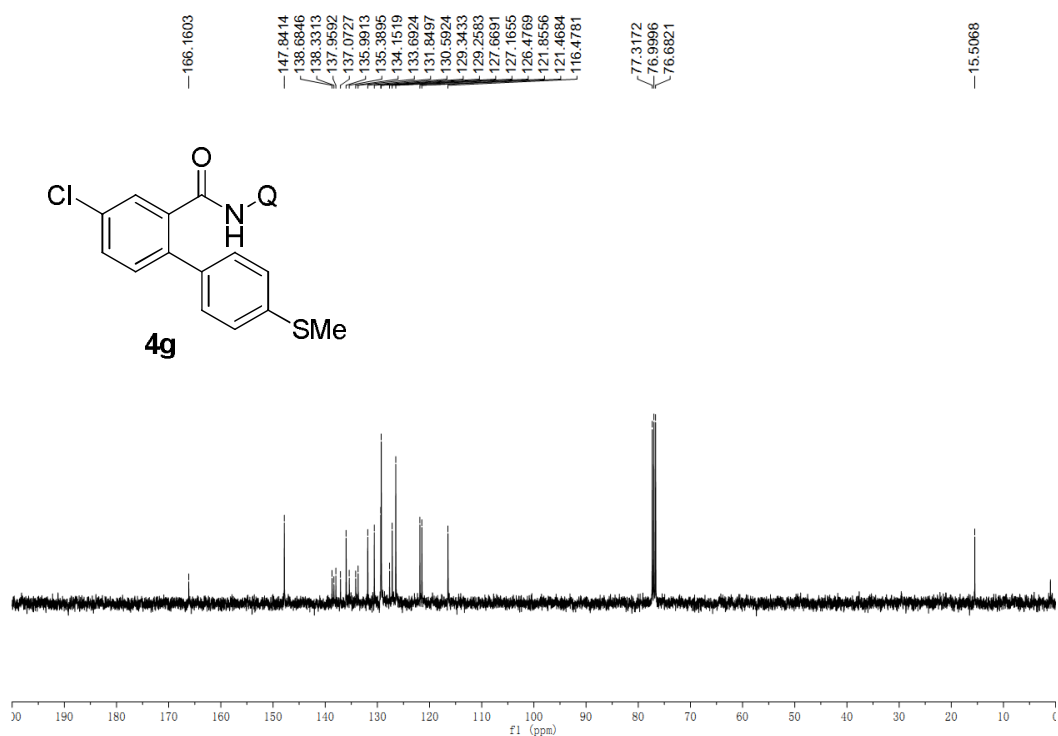
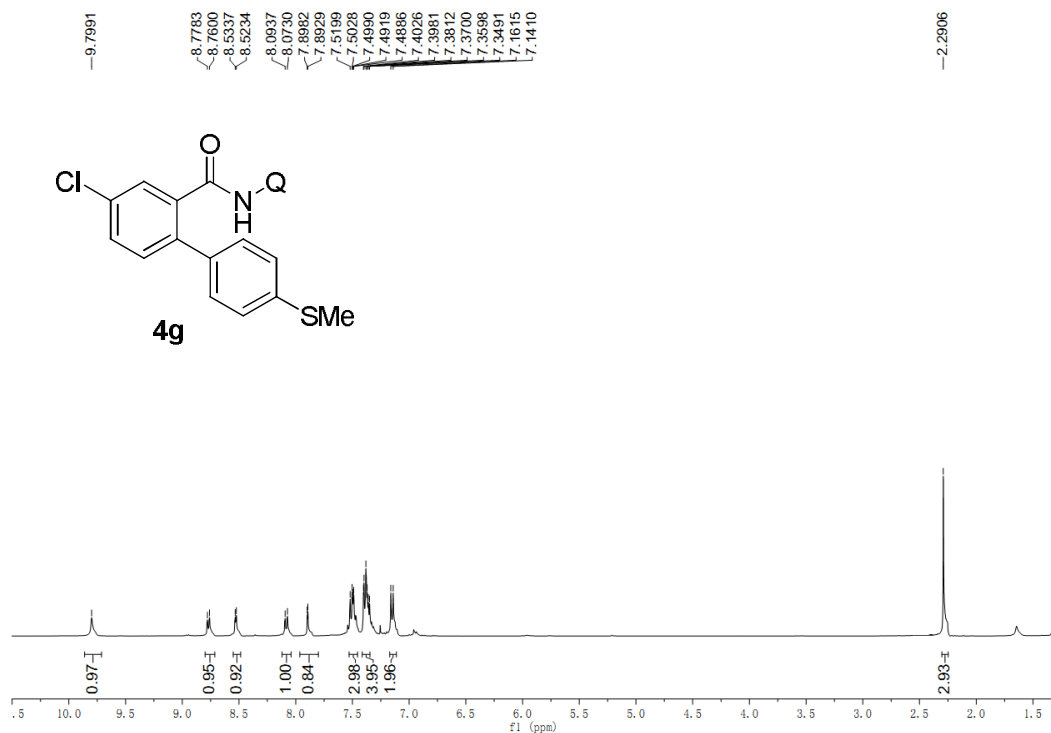


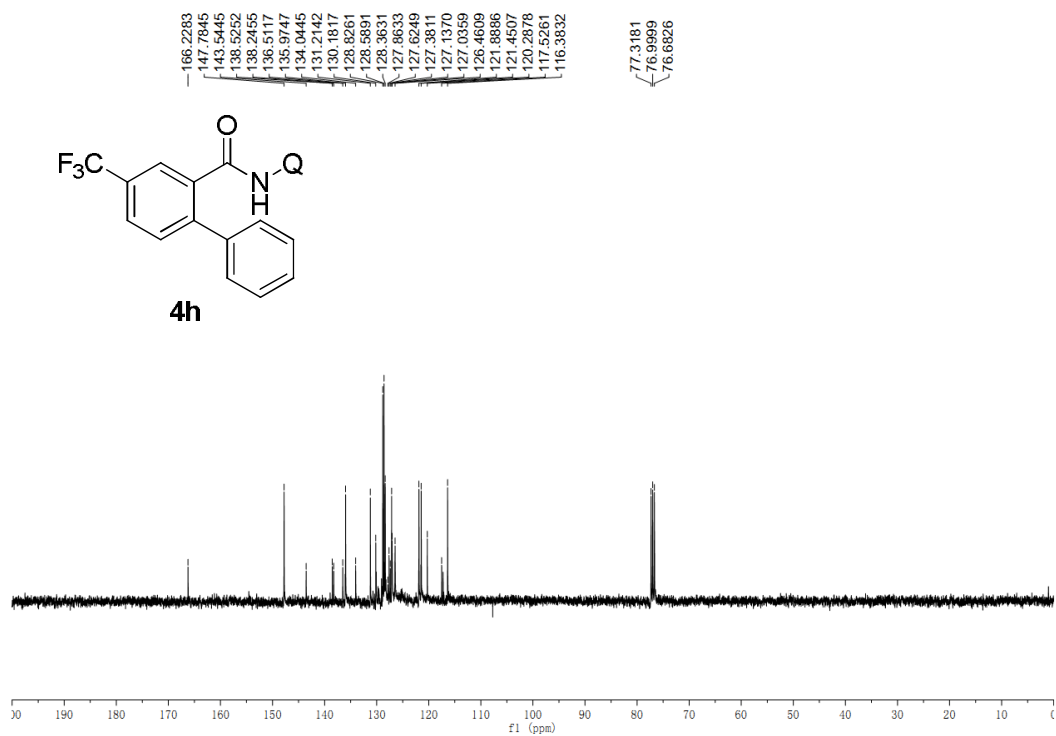
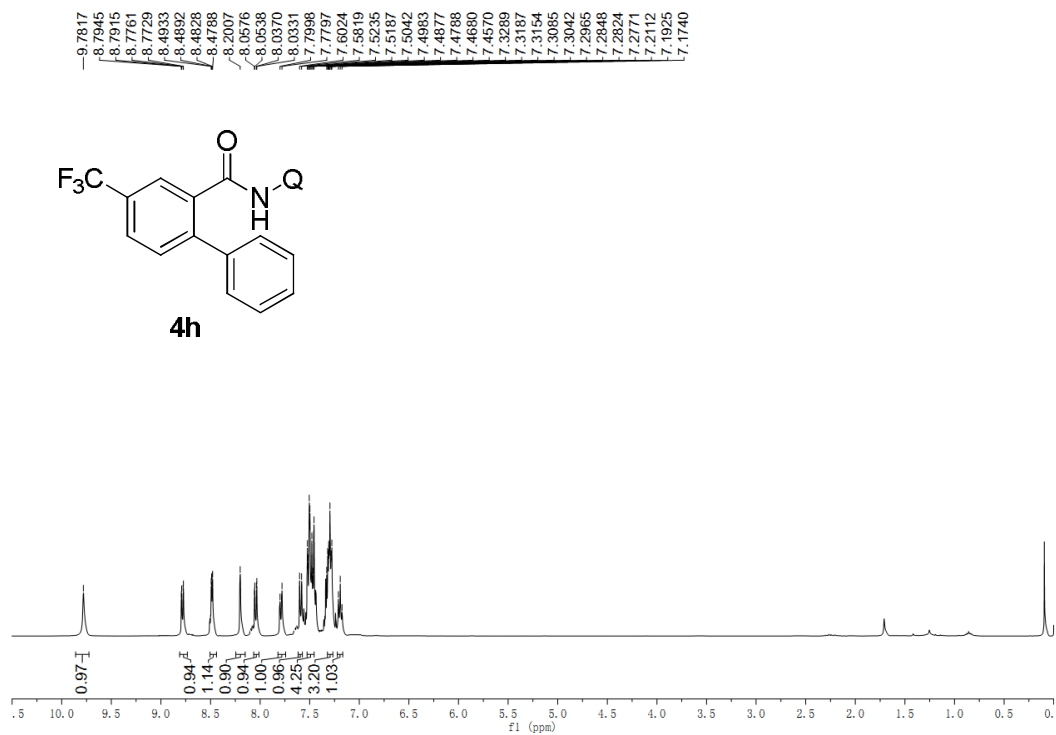
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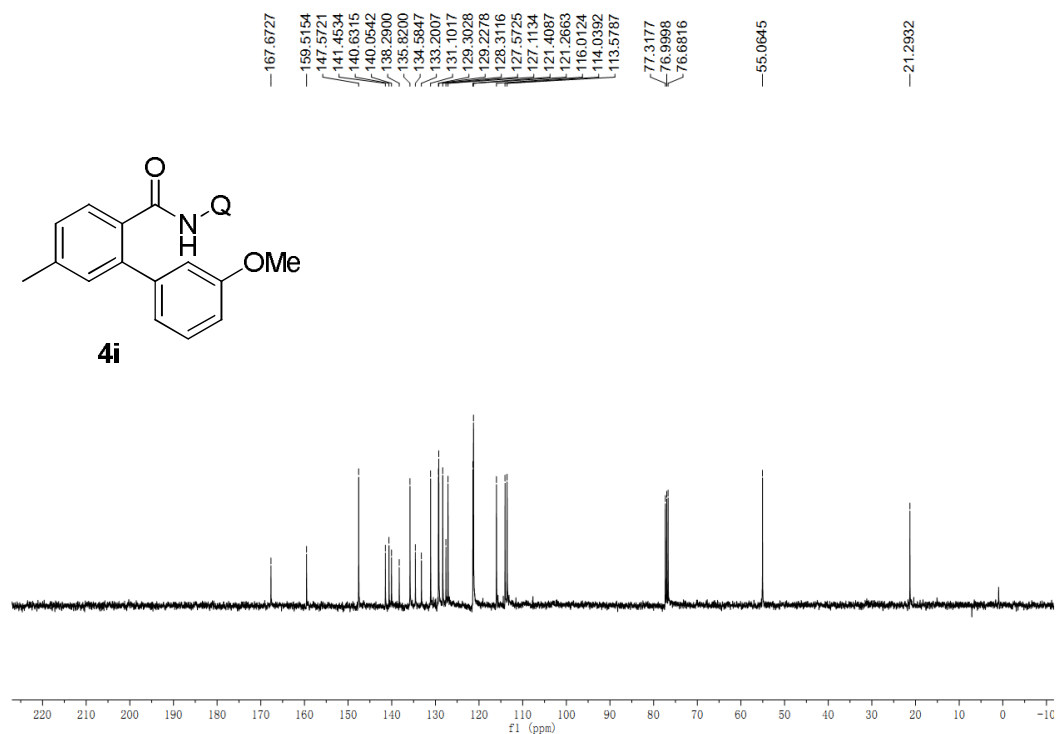
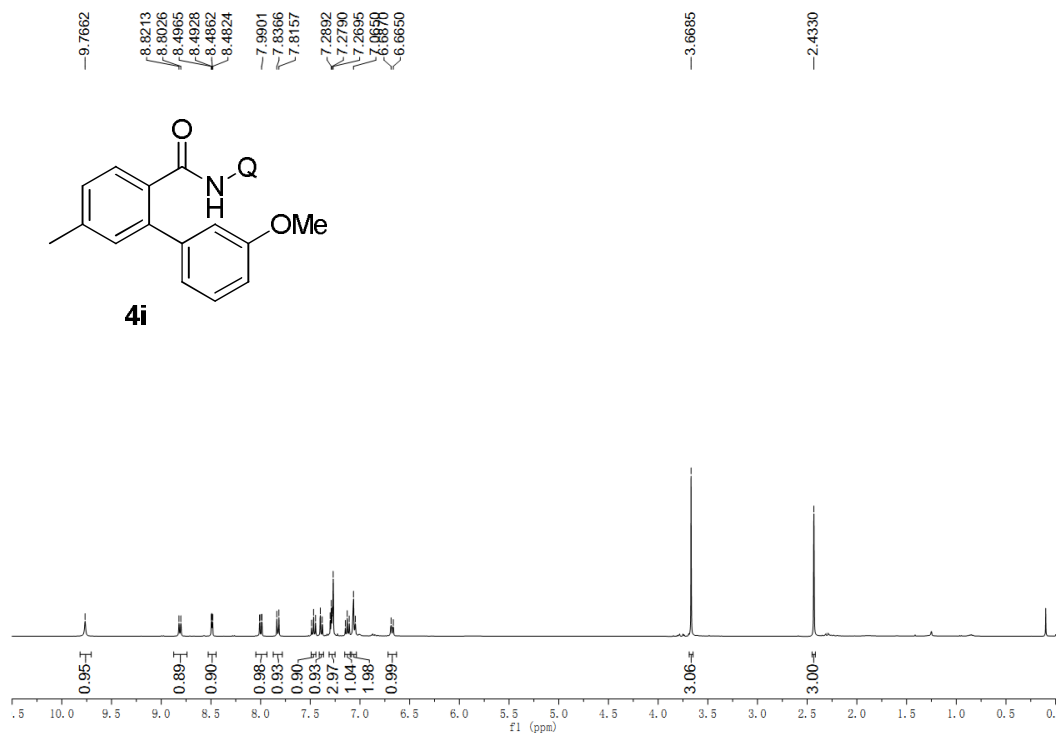


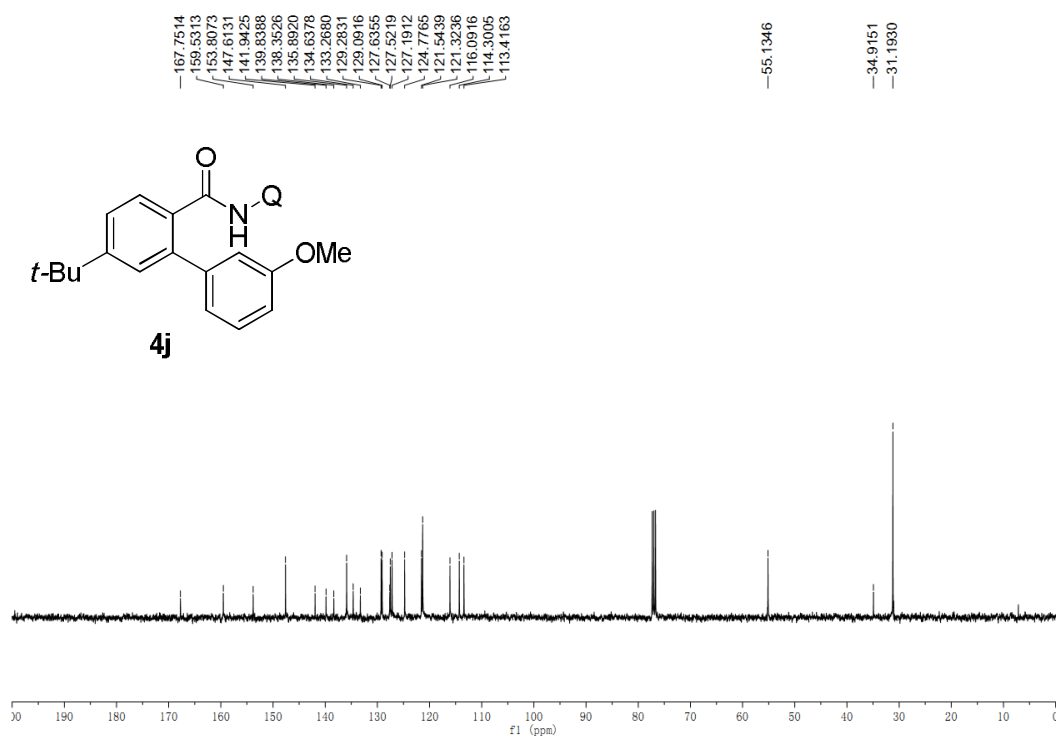
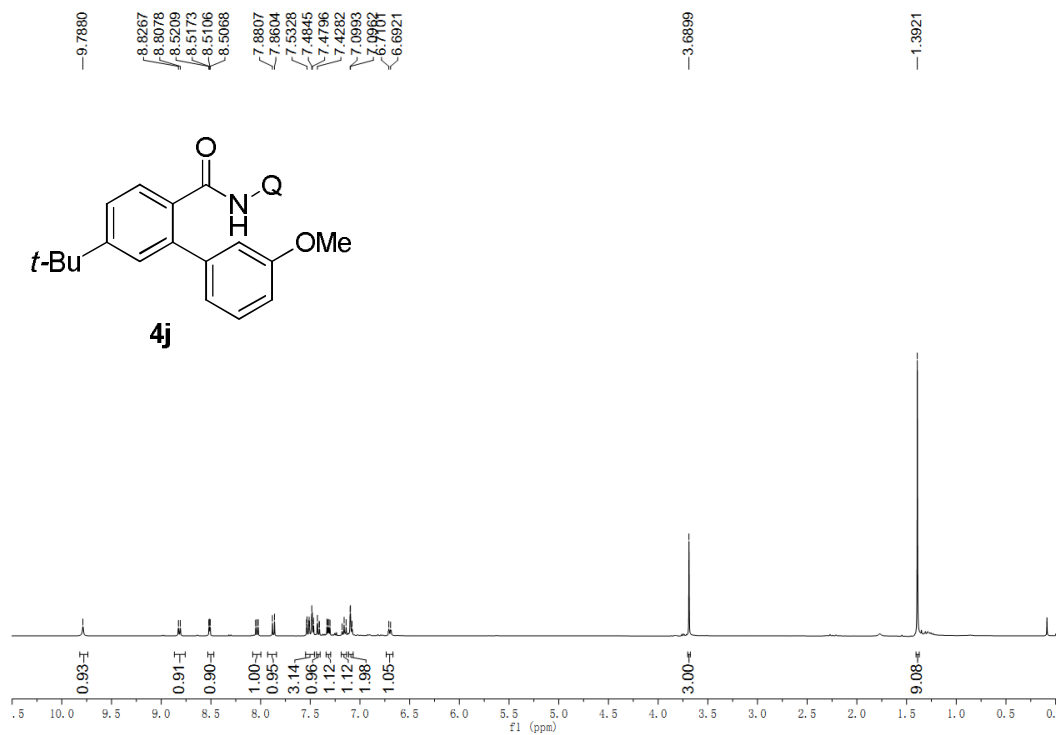
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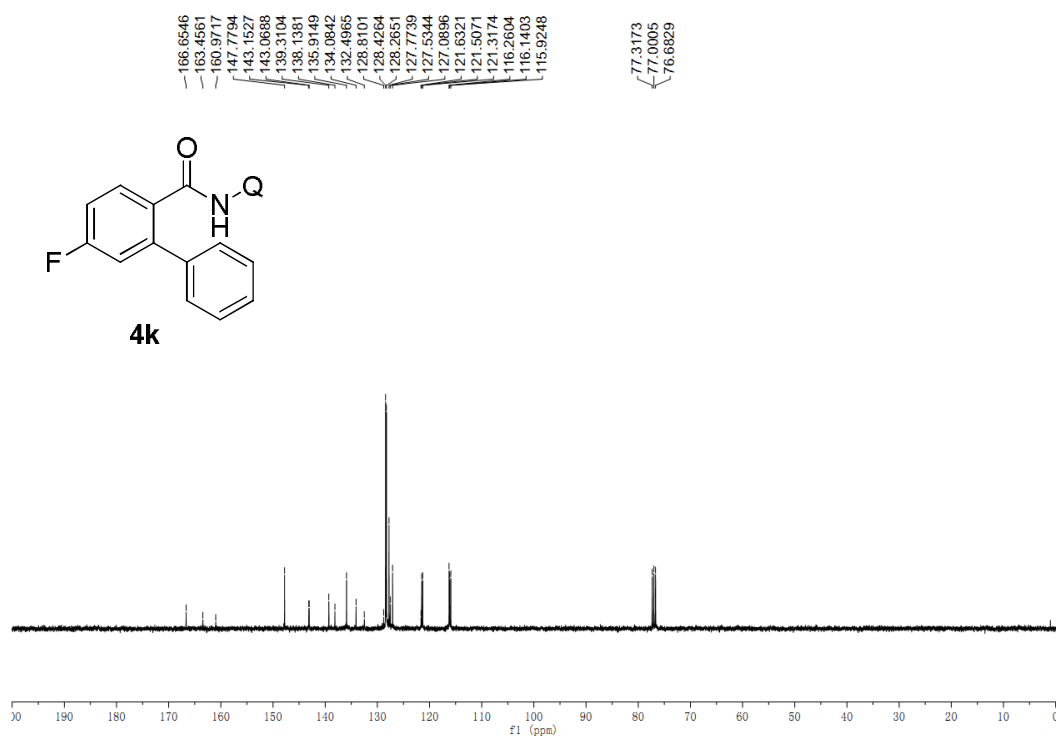
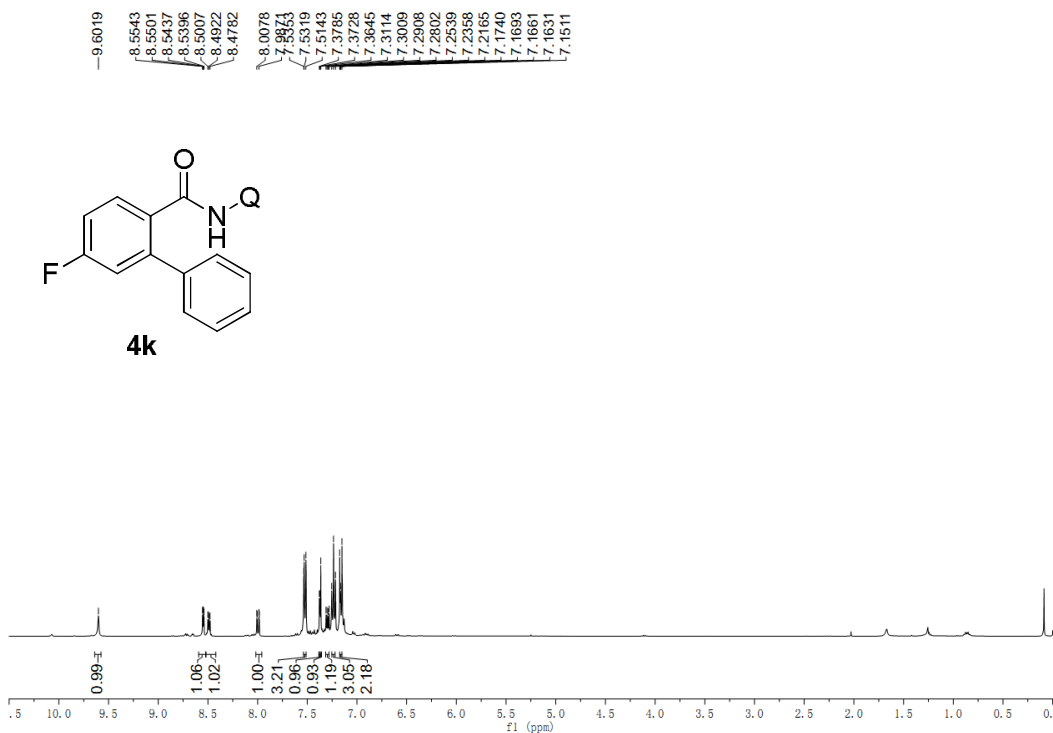


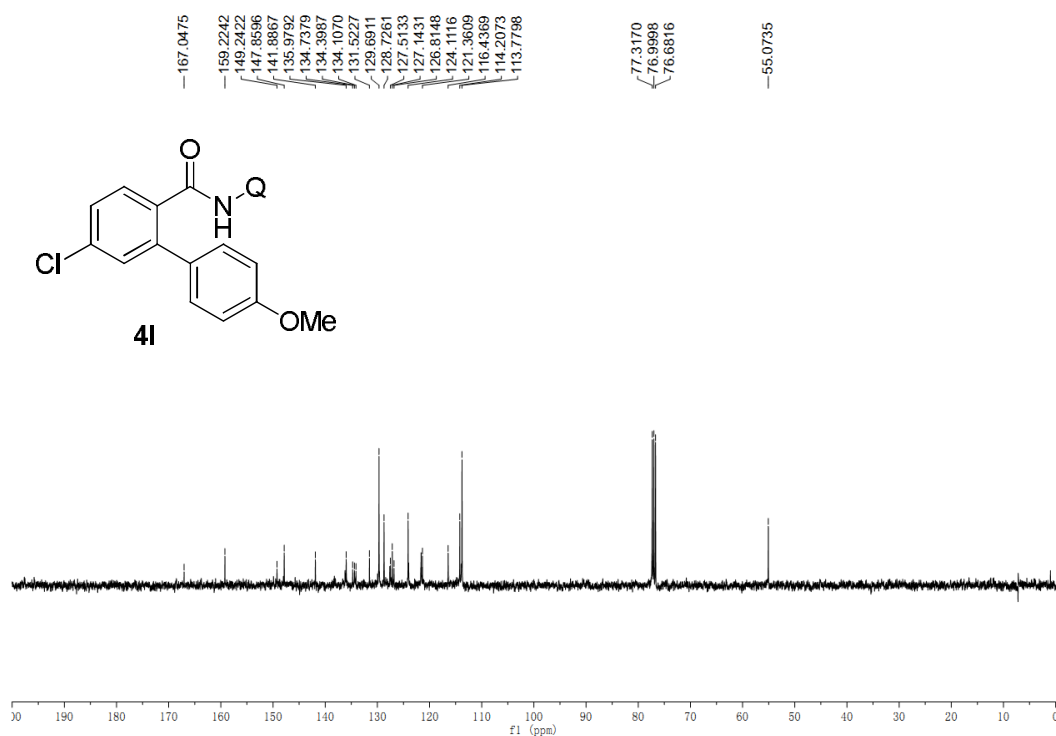
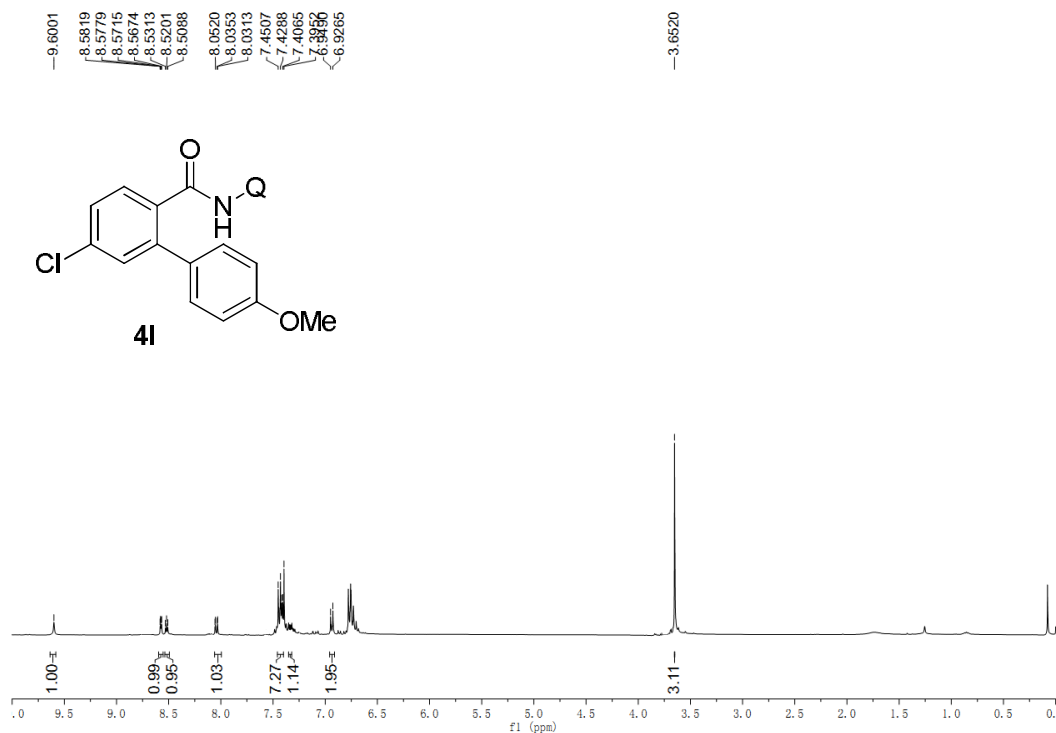


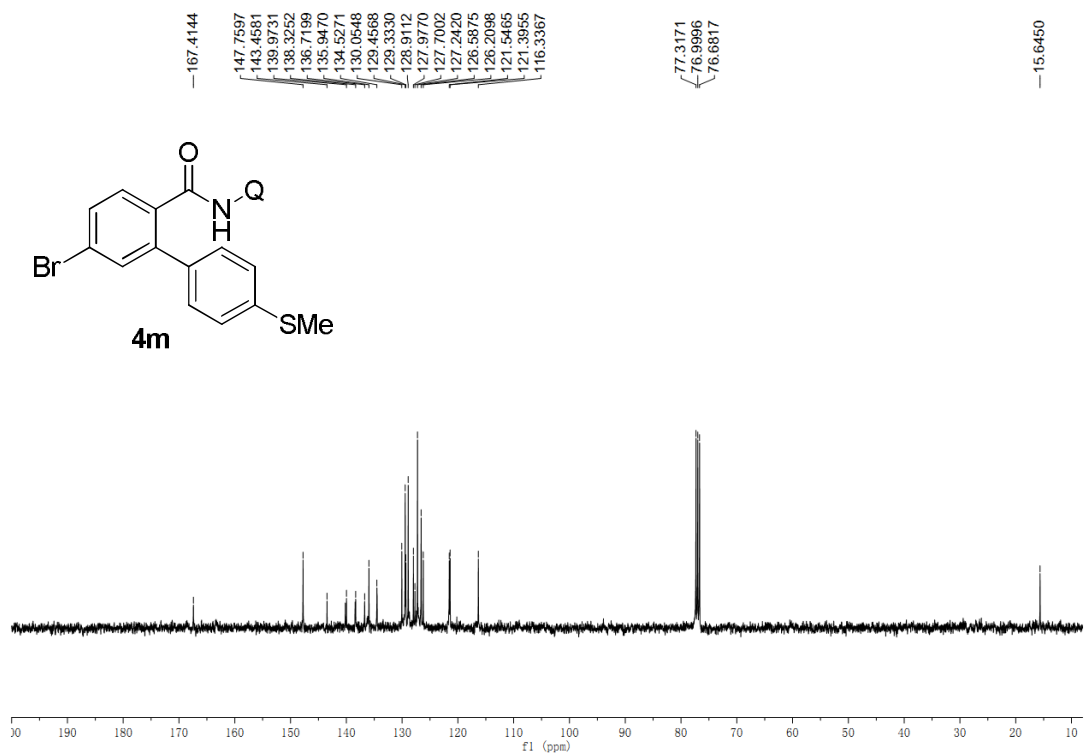
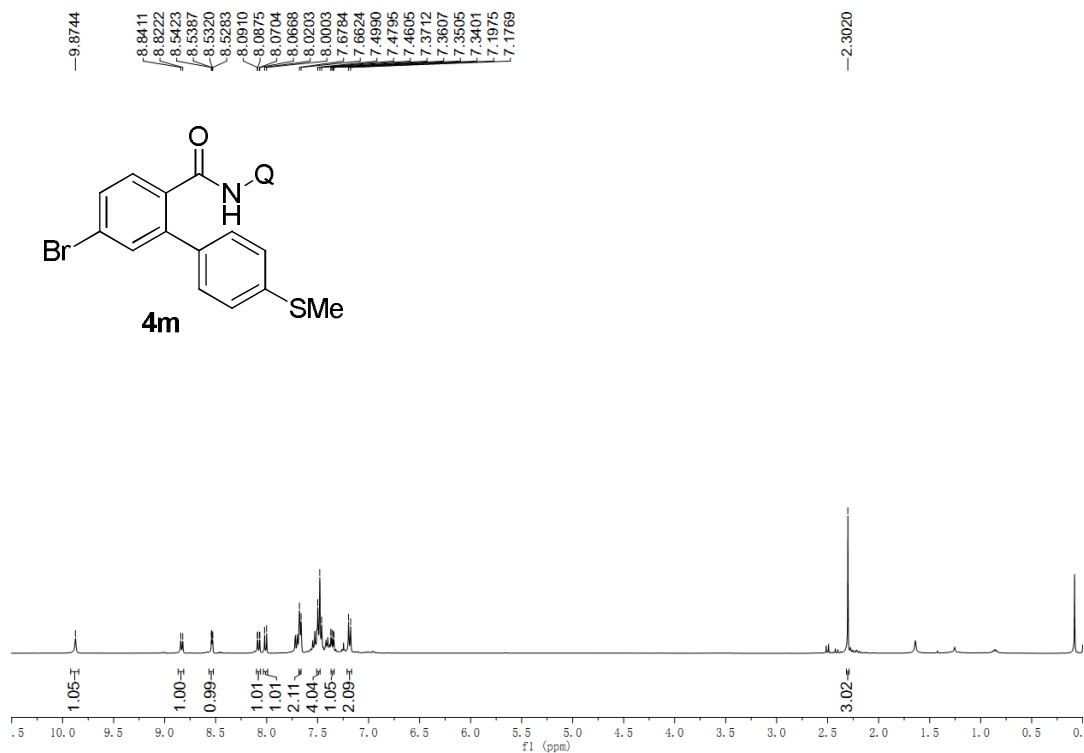


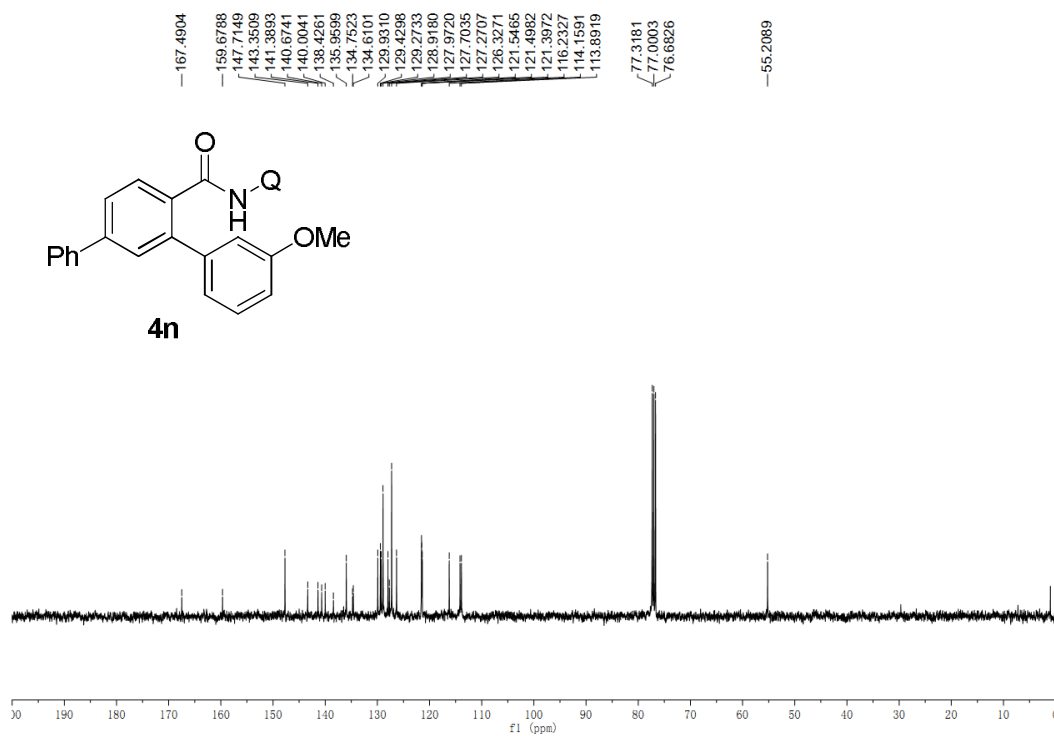
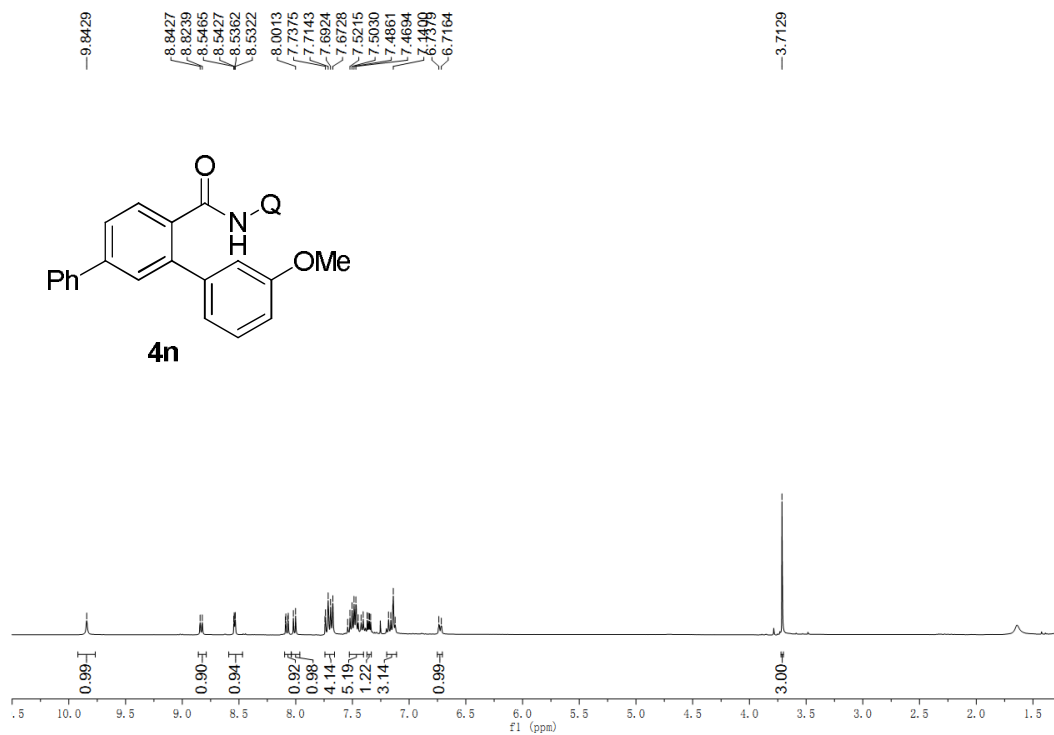


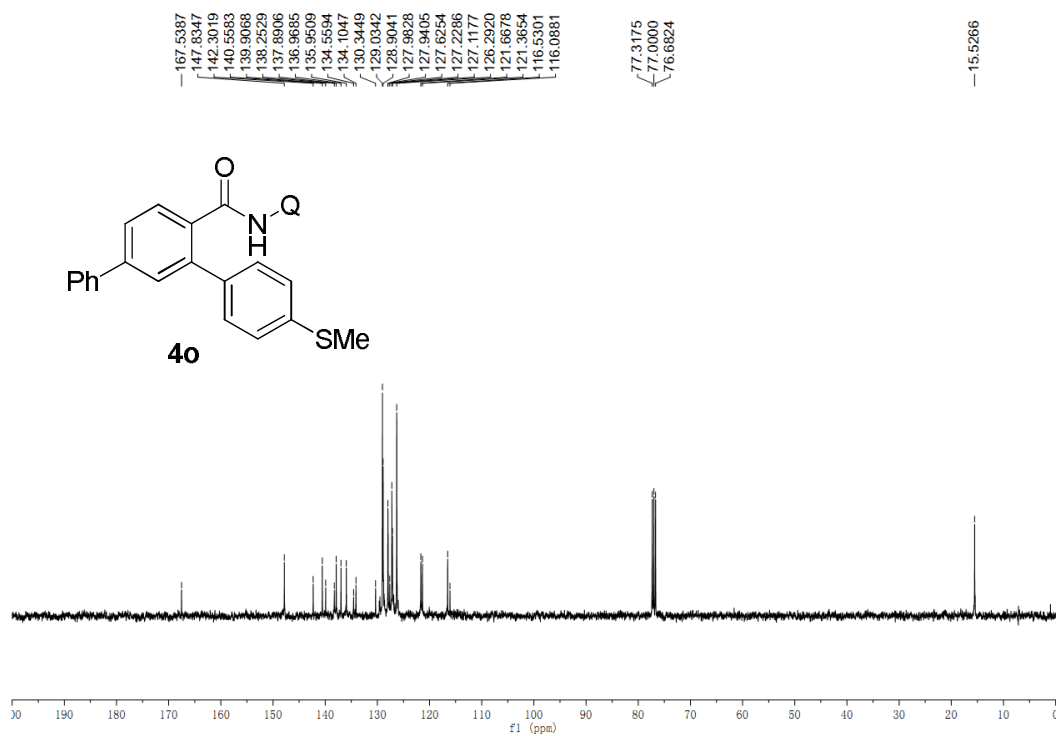
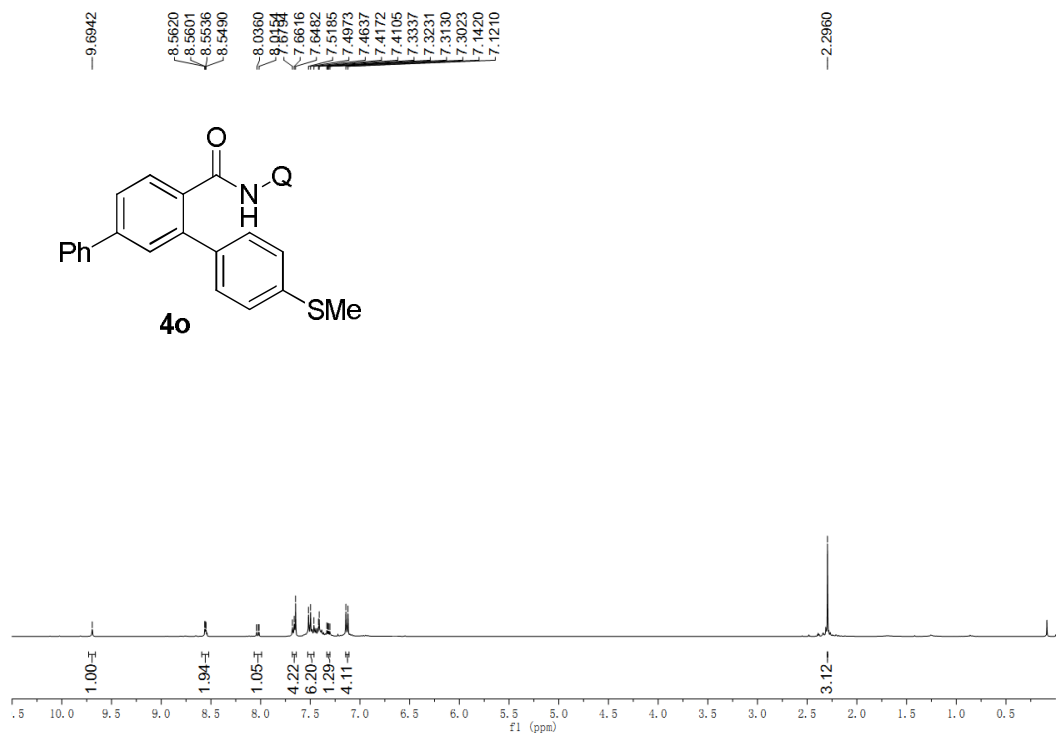


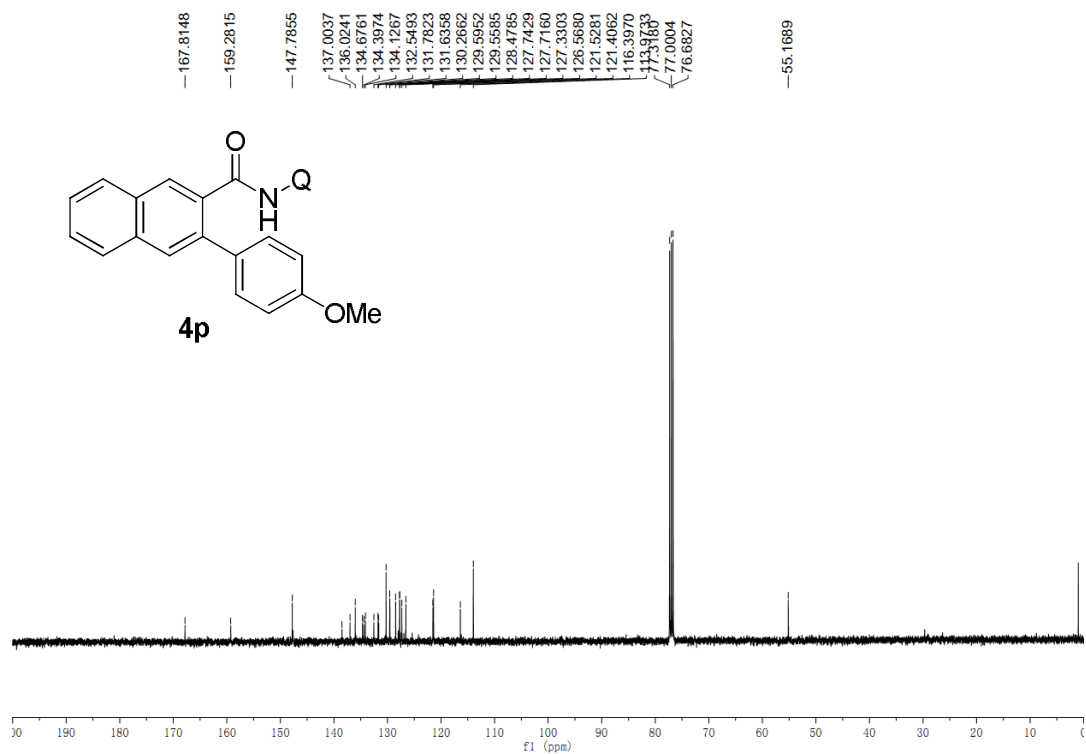
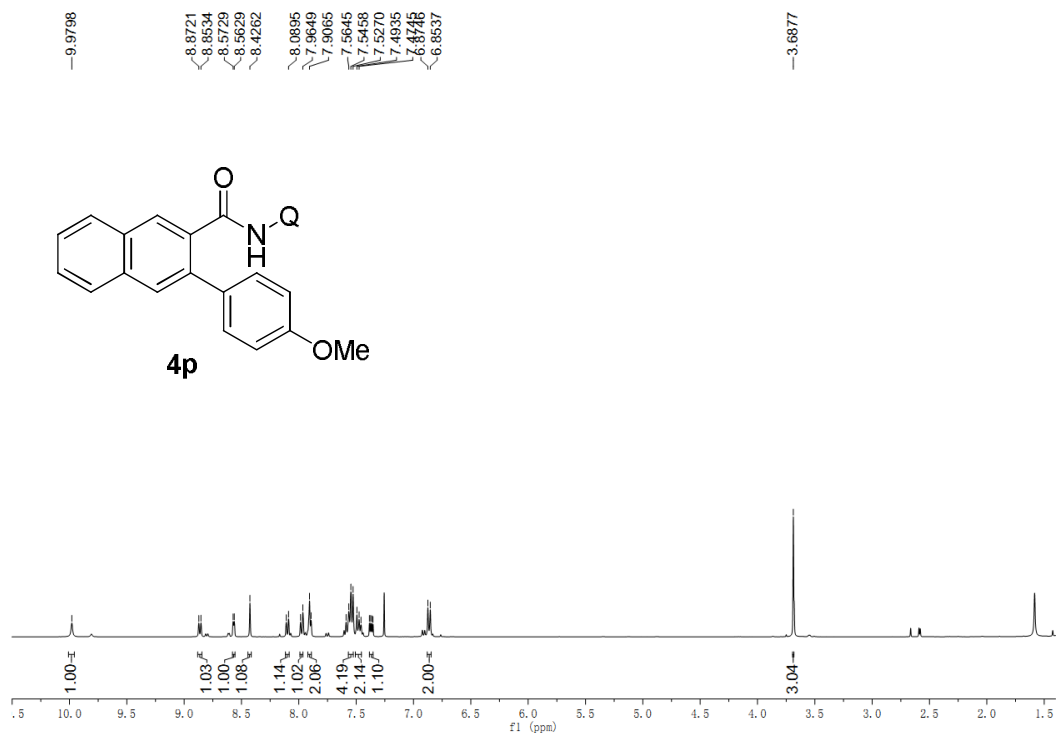


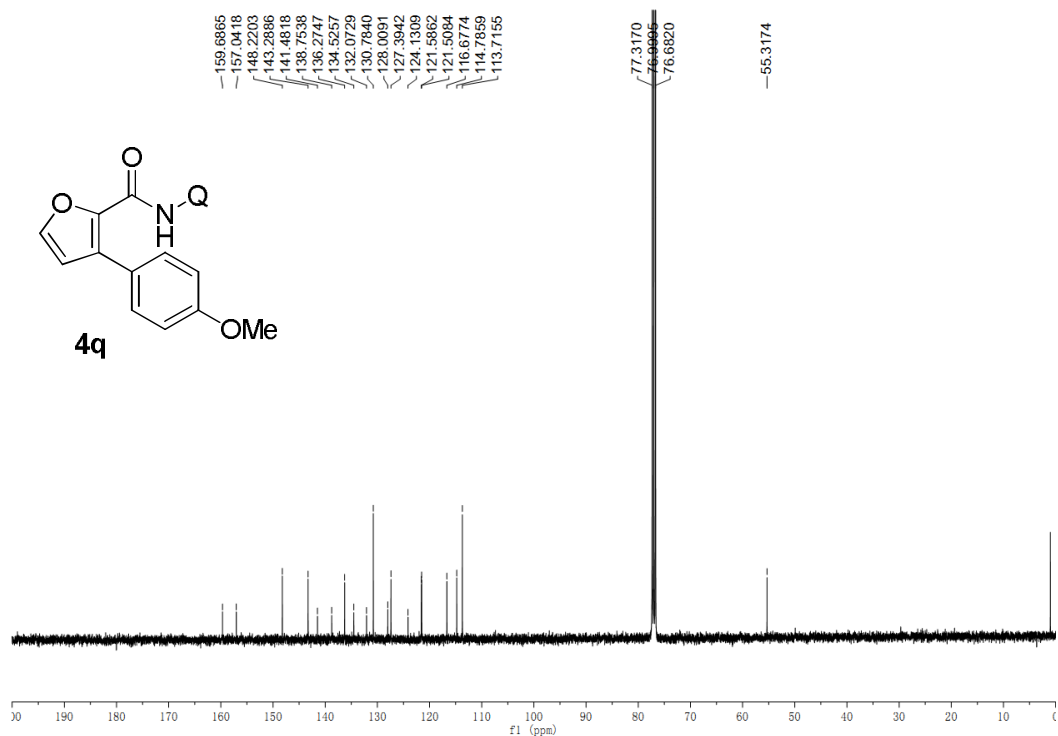
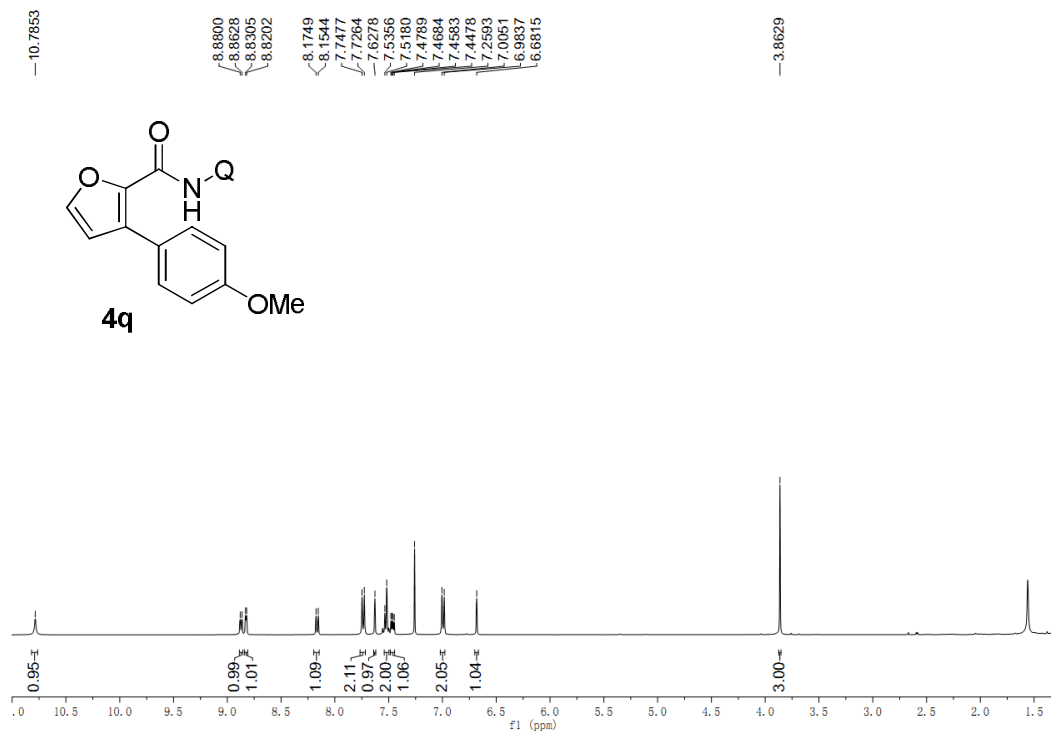


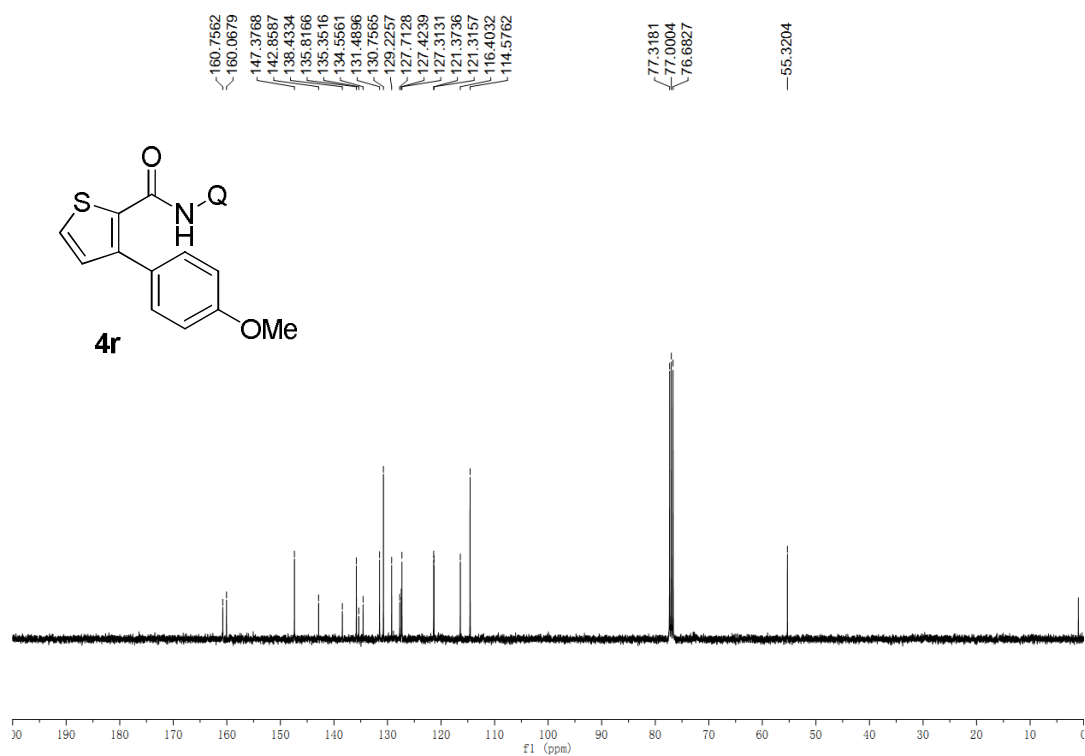
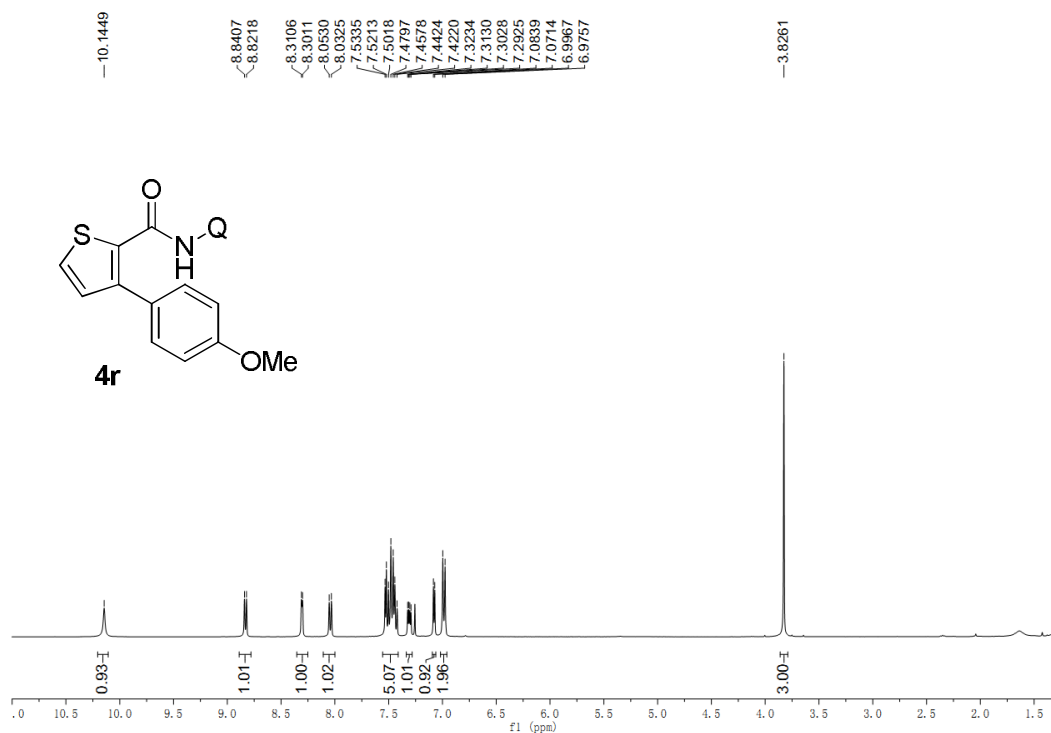


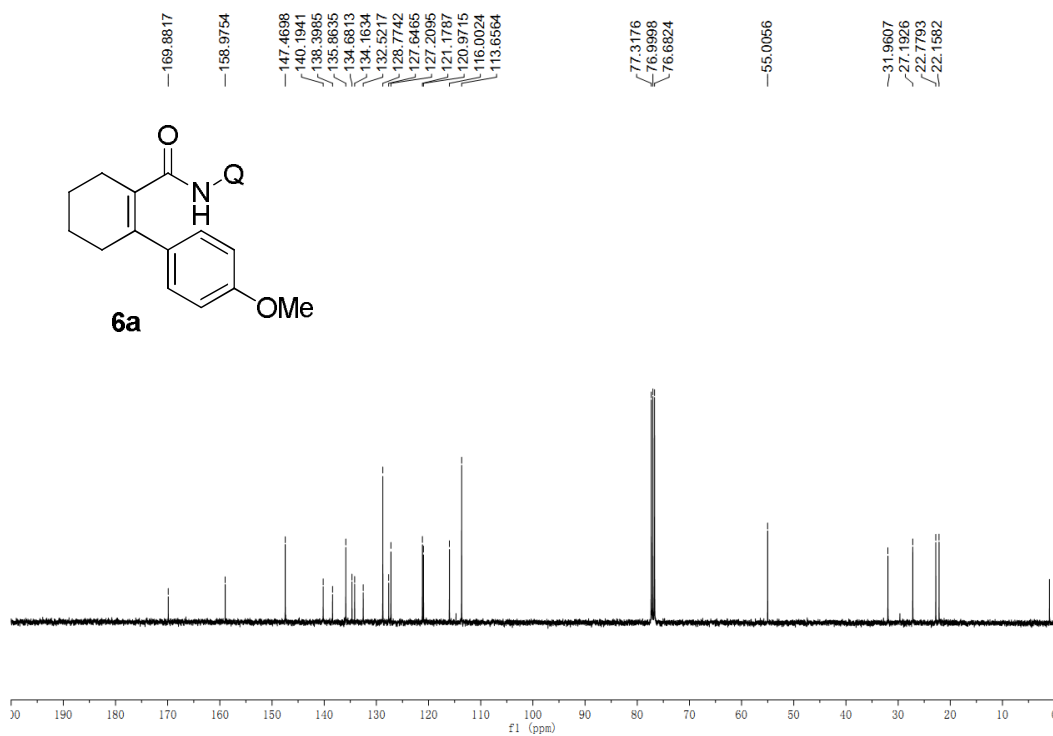
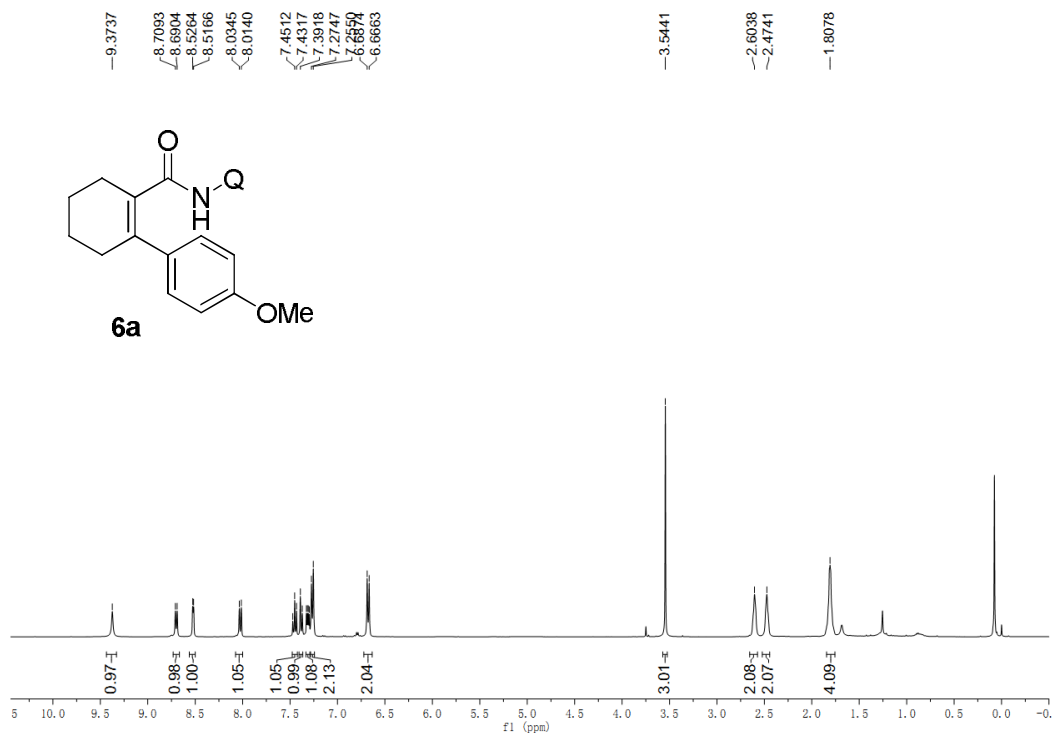


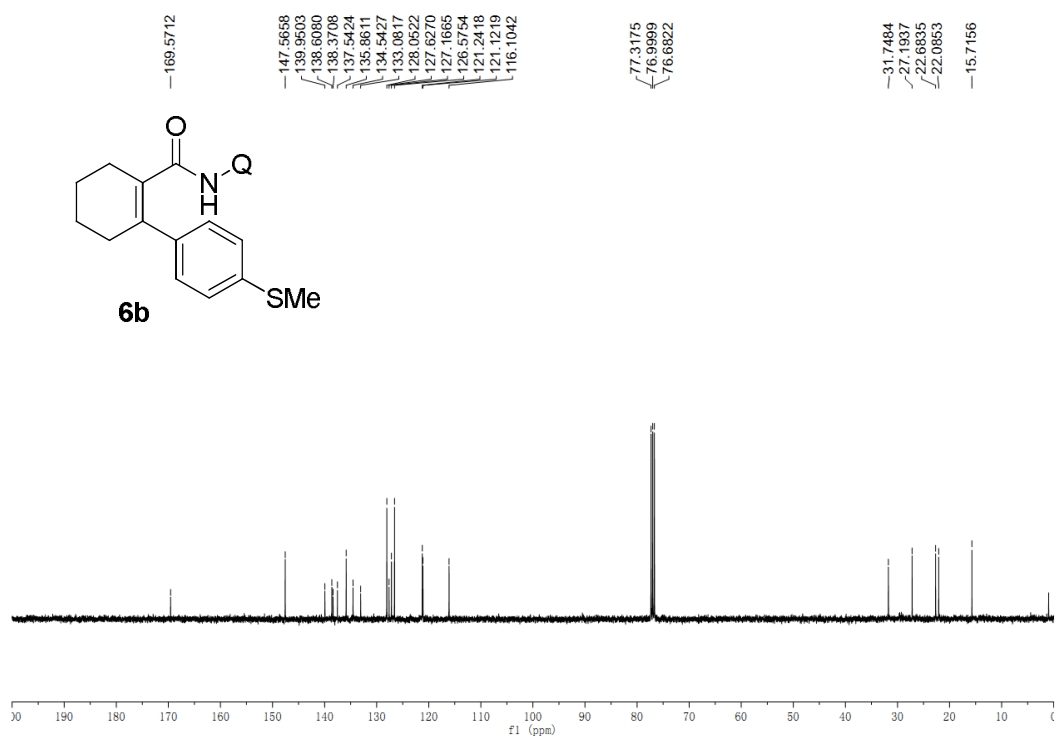
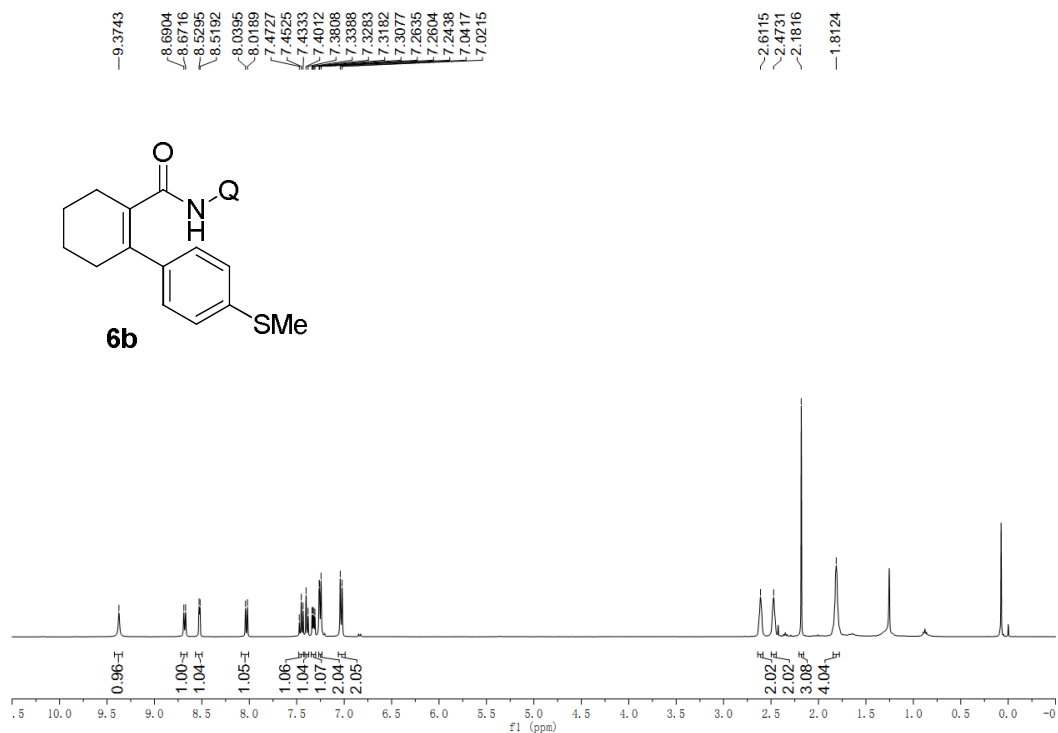


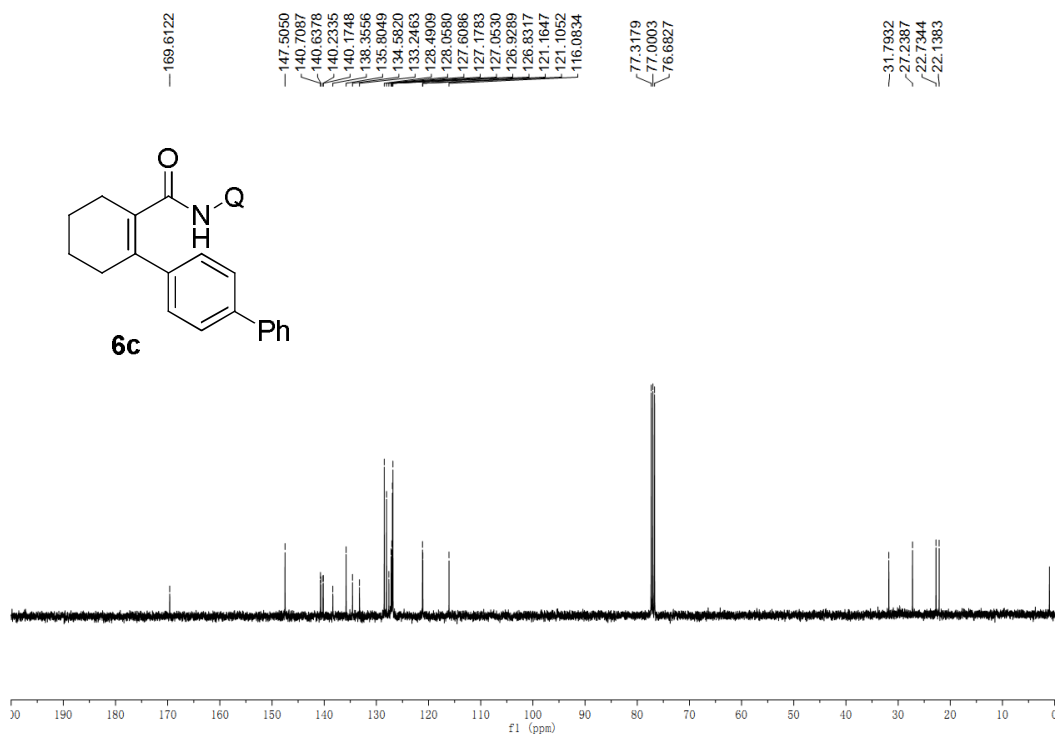
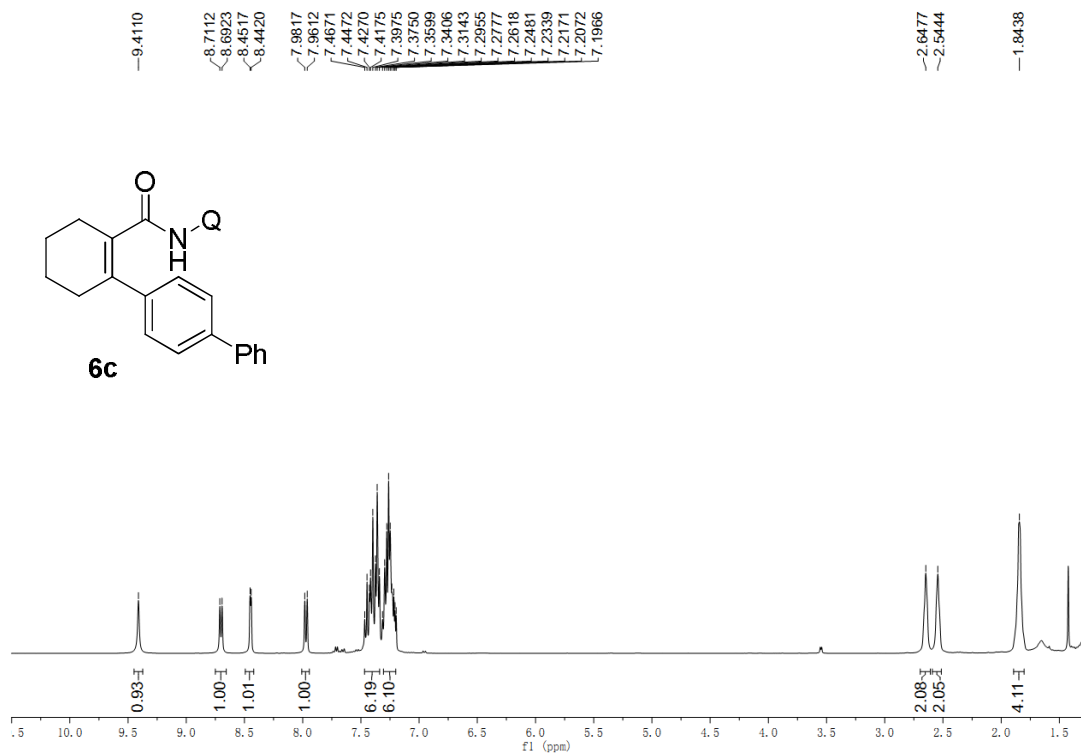




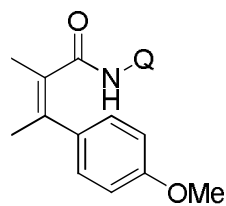






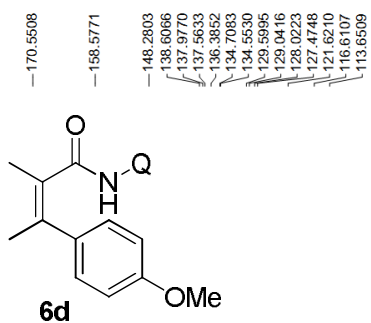
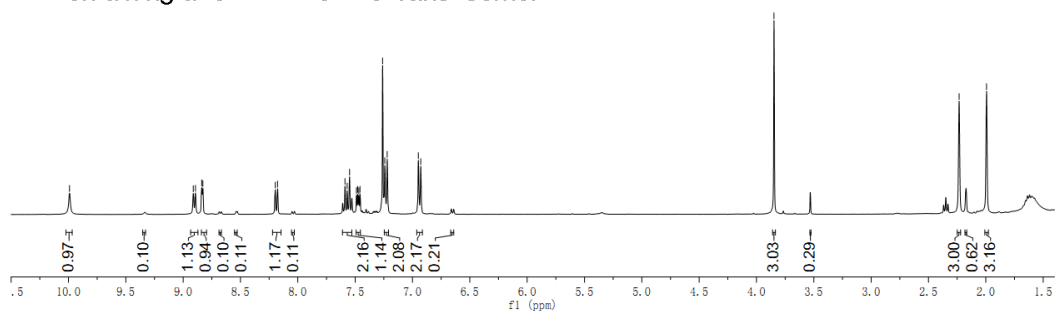


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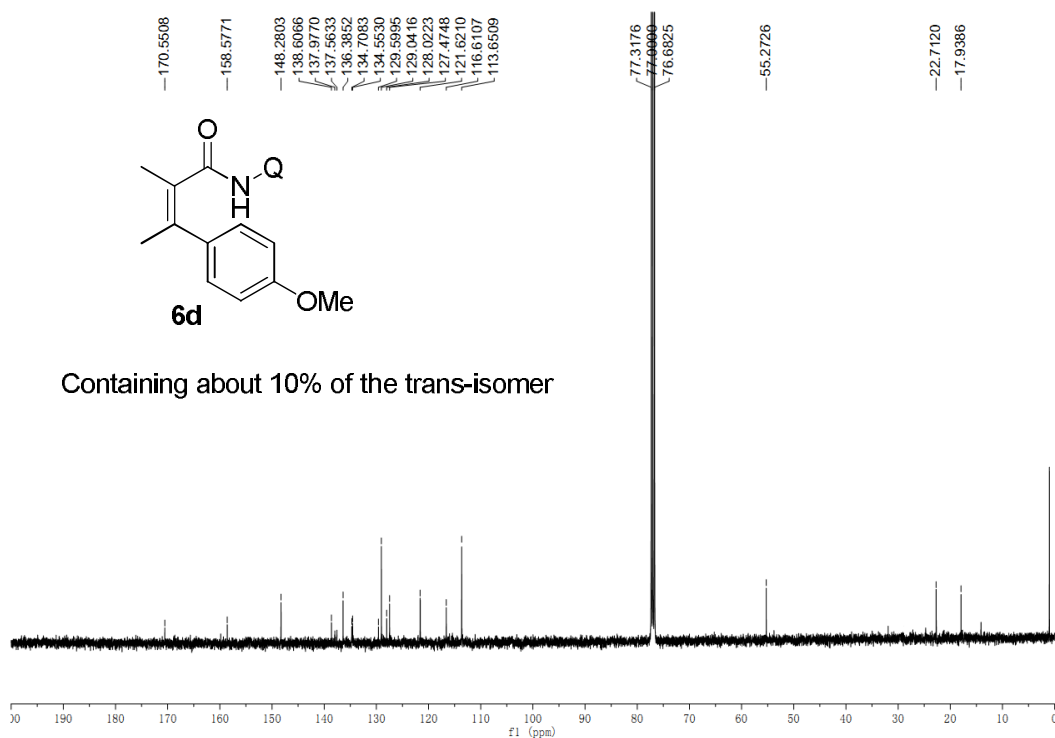
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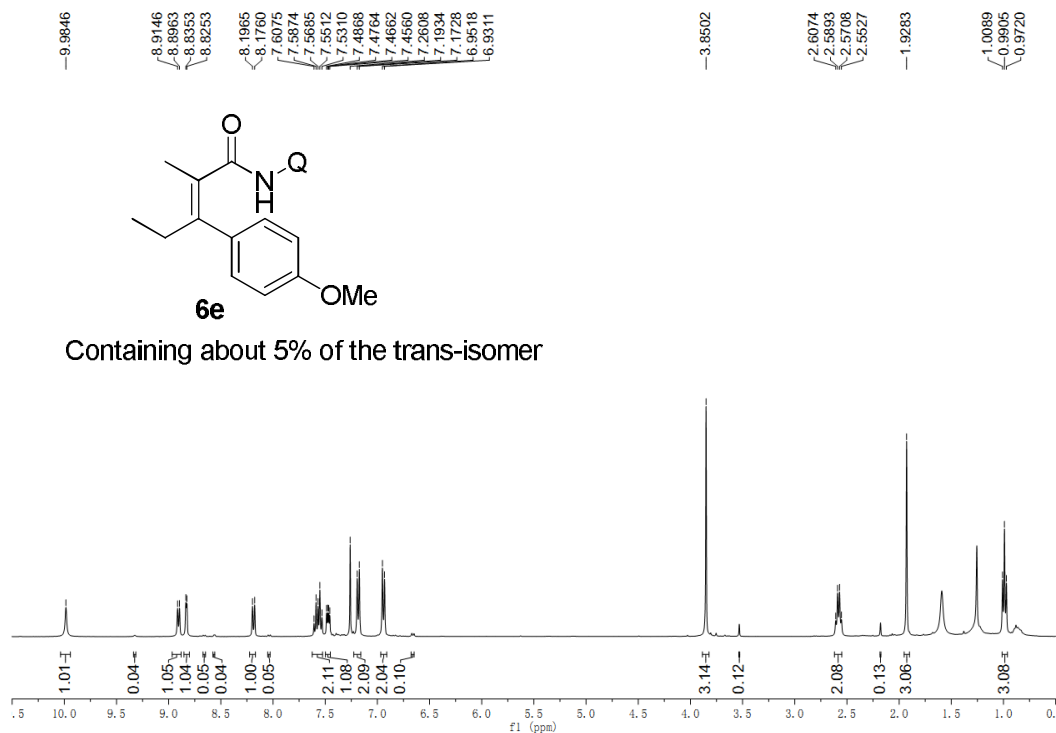
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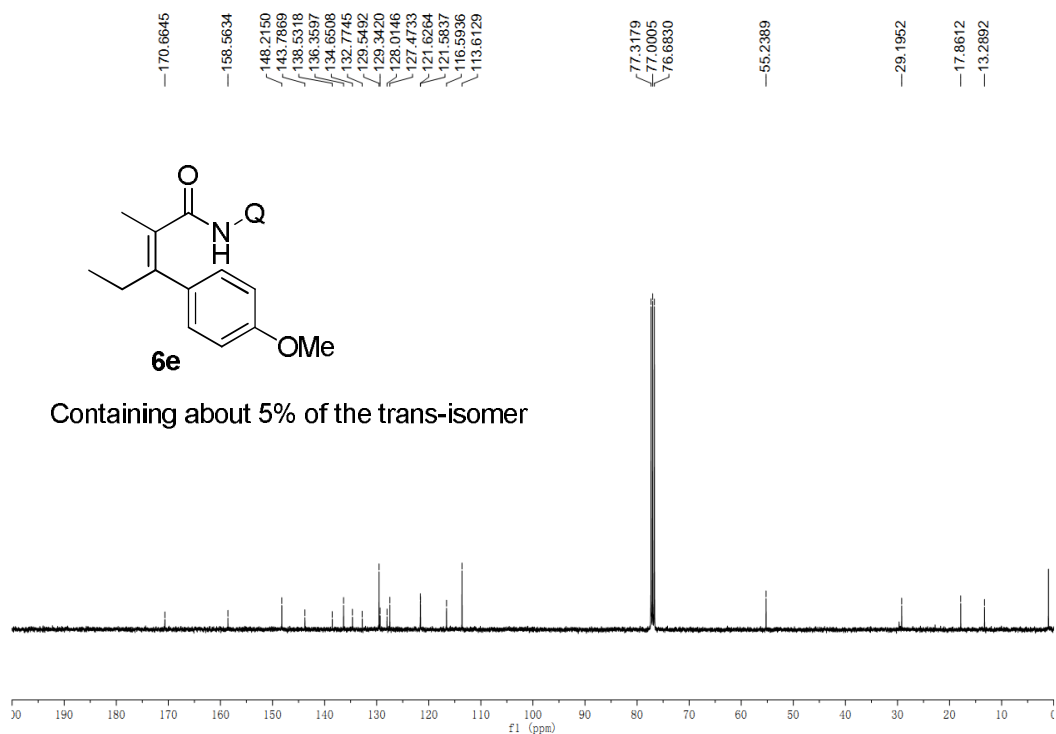
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Containing about 10% of the trans-isomer

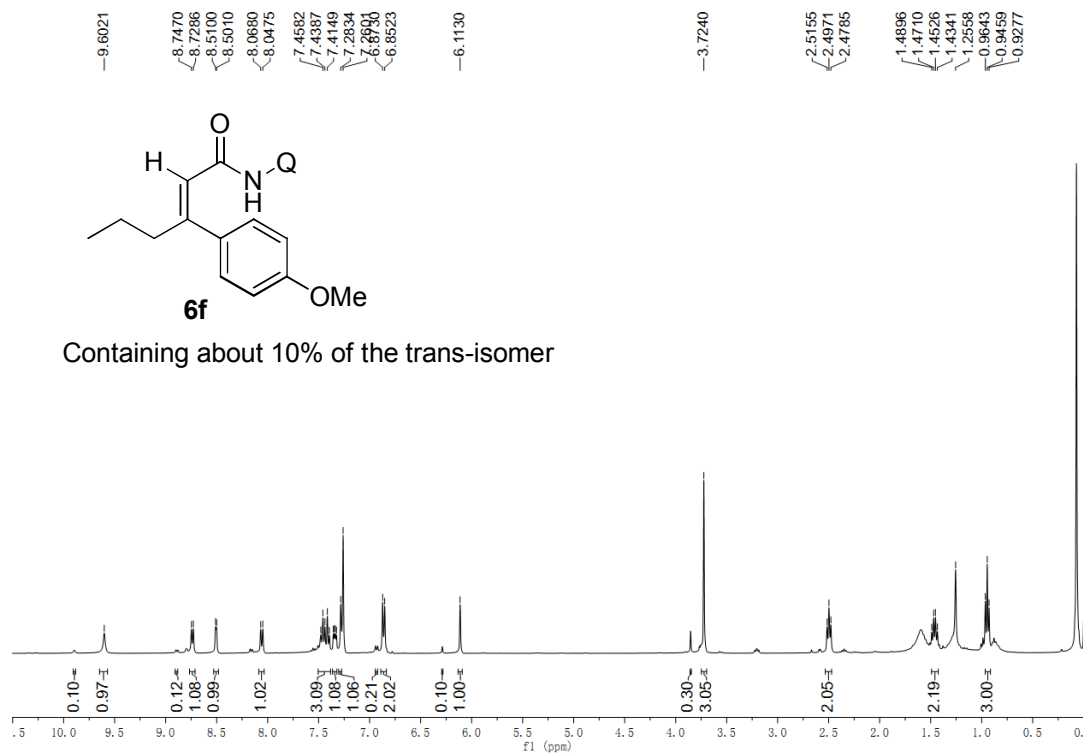




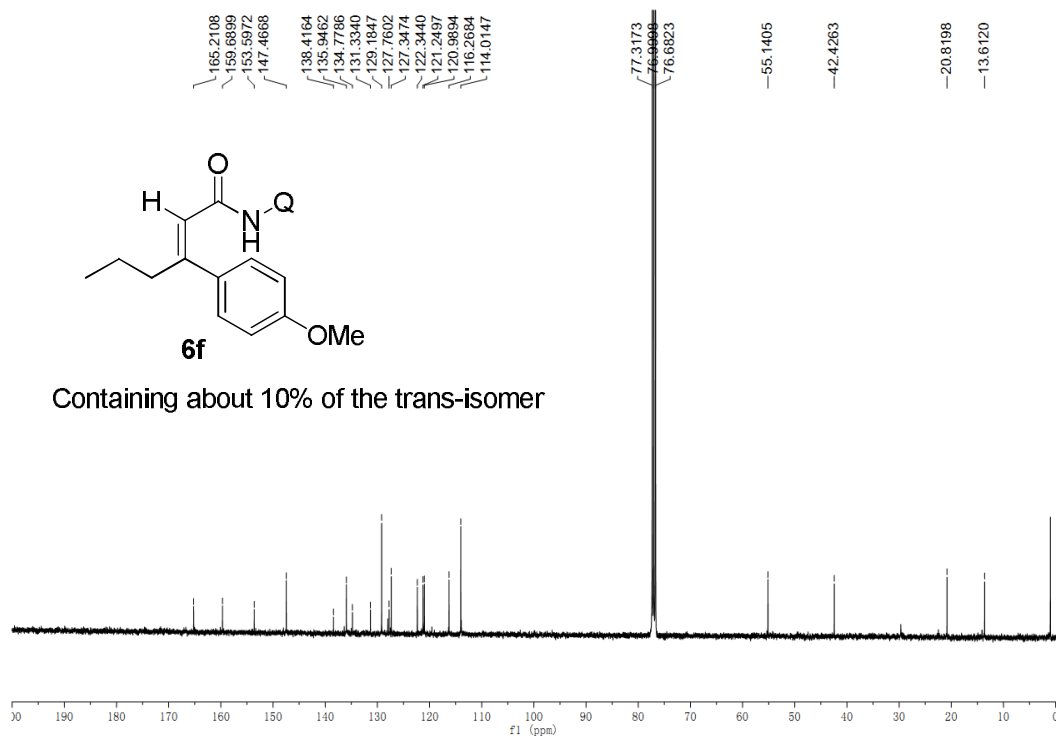
Containing about 5% of the trans-isomer



Containing about 5% of the trans-isomer



Containing about 10% of the trans-isomer



Containing about 10% of the trans-isomer

NOE of (Z)-**6f**:

