

Supporting Information

Titanium-Catalyzed Cyano-Borrowing Reaction for the Direct Amination of Cyanohydrins with Ammonia

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I. General information

^1H and ^{13}C NMR spectra were recorded on a Bruker Avance 600 MHz instruments. Chemical shifts were reported in parts per million (ppm), and the residual solvent peak was used as an internal reference: proton (chloroform δ 7.26), carbon (chloroform δ 77.0) or tetramethylsilane (TMS δ 0.00) was used as a reference. Multiplicity was indicated as follows: s (singlet), d (doublet), t (triplet), q (quartet), m (multiplet), dd (doublet of doublet), bs (broad singlet). Coupling constants were reported in Hertz (Hz). All high resolution mass spectra (**HRMS**) were obtained on a Bruker Apex-2. For thin layer chromatography (**TLC**), Qingdao Haiyang Chemical were used, and compounds were visualized with a UV light at 254 nm. Further visualization was achieved by staining with iodine, or potassium permanganate solution followed by heating using a heat gun. Flash chromatography separations were performed on Qingdao Haiyang Chemical 300-400 mesh silica gel. All commercially available reagents were used as received for the reactions without any purification. All solvents were dried on alumina columns using a solvent dispensing system.

II. General procedure for the Titanium-Catalyzed Cyano-Borrowing Reaction for the Direct Amination of Cyanohydrins with Ammonia.

Method A: To a vial equipped with a dried stir bar was added ammonia (7.0 M solution in MeOH) 0.2 mL, ketone cyanohydrins (0.4 mmol), $\text{Ti}(\text{O}^i\text{Pr})_4$ (10 mol%), benzoic acid (40 mol%), TBME (0.8 mL) in the glovebox. The reaction mixture was taken outside the glovebox after stirring and mixed evenly. Then, the reaction mixture was allowed to stir at the settle temperature for 18 hours. The reaction mixture was added to water (10 mL), extracted with EtOAc (3×5 mL). The organic layer was washed with aqueous NaHCO_3 and brine and dried over Na_2SO_4 . And the residue was purified by column chromatography with silica gel to give pure products. All ketone cyanohydrins used method A.

Method B: To a vial equipped with a dried stir bar was added 0.2 mL ammonia (7.0 M solution in MeOH), aldehyde cyanohydrins (0.4 mmol), $\text{Ti}(\text{O}^i\text{Pr})_4$ (10 mol%), MeOH (0.8 mL) in the glovebox. The reaction mixture was taken outside the glovebox after stirring and mixed evenly. Then, the reaction mixture was allowed to stir at the settle temperature for 18 hours. The reaction mixture was added to water (10 mL), extracted with EtOAc (3×5 mL). The organic layer was washed with aqueous NaHCO_3 and brine and dried over Na_2SO_4 . And the residue was purified by column chromatography with silica gel to give pure products. All aldehyde cyanohydrins used method B.

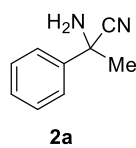
III. General Procedure for the Gram Scale reaction.

Scheme 5(1): To a vial equipped with a dried stir bar was added ammonia (7.0 M solution in MeOH) 5.0 mL, ketone cyanohydrins (10.0 mmol), $\text{Ti}(\text{O}^i\text{Pr})_4$ (10 mol%), benzoic acid (40 mol%), TBME (15 mL) in the glovebox. The reaction mixture was taken outside the glovebox after stirring and mixed evenly. Then, the reaction mixture was allowed to stir at the settle temperature for 18 hours. The reaction mixture was added to water (30 mL), extracted with EtOAc (3×25 mL). The organic layer was washed with aqueous NaHCO_3 and brine and dried over Na_2SO_4 . And the residue was purified by column chromatography with silica gel to give 0.83 g of product **2a** with 57% isolated yield.

Scheme 5(2): To a vial equipped with a dried stir bar was added 10.0 mL ammonia (7.0 M solution in MeOH), aldehyde cyanohydrins (20 mmol), $\text{Ti}(\text{O}^i\text{Pr})_4$ (10 mol%), MeOH (20 mL) in the glovebox. The reaction mixture was taken outside the glovebox after stirring and mixed evenly. Then, the reaction mixture was allowed to stir at the settle temperature for 18 hours. The reaction mixture was added to water (30 mL), extracted with EtOAc (3×25 mL). The organic layer was washed with aqueous NaHCO_3 and brine and dried over Na_2SO_4 . And the residue was purified by column chromatography with silica gel to give 2.30 g of product **6a** with 87% isolated yield.

IV The analytical and spectral characterization data

2-amino-2-phenylpropanenitrile (**2a**)^[1]

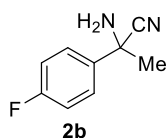


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.4$ in hexane:ethyl acetate = 2:1) resulting in 37.4 mg of Colorless oil product **2a** with 64% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.60 (d, $J = 7.6$ Hz, 2H), 7.34 (t, $J = 7.5$ Hz, 2H), 7.30 (d, $J = 7.3$ Hz, 1H), 2.27 (s, 2H), 1.73 (s, 3H).

^{13}C NMR (151 MHz, CDCl_3) δ 141.23, 128.85, 128.65, 124.94, 124.17, 53.64, 31.79.

2-amino-2-(4-fluorophenyl)propanenitrile (**2b**)^[2]

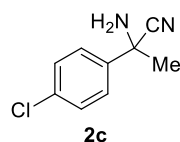


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.4$ in hexane:ethyl acetate = 2:1) resulting in 42.6 mg of Colorless oil product **2b** with 65% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.57 (dd, $J = 8.2$ Hz, 5.3 Hz, 2H), 7.01 (t, $J = 8.5$ Hz, 2H), 1.99 (s, 2H), 1.67 (s, 3H).

^{13}C NMR (151 MHz, CDCl_3) δ 163.53, 161.89, 137.08, 126.88 (d, $J = 8.3$ Hz), 123.97, 115.70 (d, $J = 21.8$ Hz), 53.14, 31.92.

2-amino-2-(4-chlorophenyl)propanenitrile (**2c**)^[3]

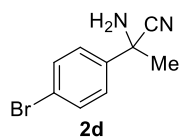


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.4$ in hexane:ethyl acetate = 2:1) resulting in 46.1 mg of Colorless oil product **2c** with 64% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.62 (d, $J = 8.6$ Hz, 2H), 7.39 (d, $J = 8.6$ Hz, 2H), 2.15 (s, 1H), 1.76 (s, 3H).

^{13}C NMR (151 MHz, CDCl_3) δ 139.86, 134.59, 128.99, 126.48, 123.76, 53.22, 31.81.

2-amino-2-(4-bromophenyl)propanenitrile (**2d**)^[2]

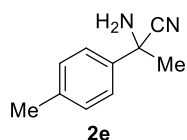


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) (R_f = 0.4 in hexane:ethyl acetate = 2:1) resulting in 57.3 mg of yellow oil product **2d** with 65% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.74-7.44 (m, 4H), 2.18 (s, 2H), 1.77 (s, 3H).

^{13}C NMR (151 MHz, CDCl_3) δ 140.40, 131.97, 126.80, 123.68, 122.71, 53.28, 31.78.

2-amino-2-(p-tolyl)propanenitrile (**2e**)^[3]

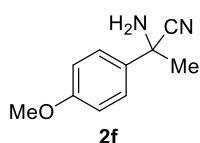


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) (R_f = 0.4 in hexane:ethyl acetate = 2:1) resulting in 39.7 mg of Colorless oil product **2e** with 62% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.46 (d, J = 8.2 Hz, 2H), 7.13 (d, J = 7.9 Hz, 2H), 2.29 (s, 3H), 1.99 (s, 2H), 1.68 (s, 3H).

^{13}C NMR (151 MHz, CDCl_3) δ 138.51, 129.47, 128.44, 124.83, 124.29, 53.41, 31.73, 21.00.

2-amino-2-(4-methoxyphenyl)propanenitrile (**2f**)^[4]

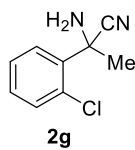


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) (R_f = 0.4 in hexane:ethyl acetate = 2:1) resulting in 42.2 mg of Colorless oil product **2f** with 60% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.49 (d, J = 8.8 Hz, 2H), 6.84 (d, J = 8.8 Hz, 2H), 3.74 (s, 3H), 1.96 (s, 2H), 1.67 (s, 3H).

^{13}C NMR (151 MHz, CDCl_3) δ 159.76, 133.33, 126.20, 124.34, 114.12, 55.36, 53.12, 31.75.

2-amino-2-(2-chlorophenyl)propanenitrile (**2g**)



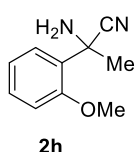
The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.4$ in hexane: ethyl acetate = 2:1) resulting in 50.4 mg of Colorless oil product **2g** with 70% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.68 (dd, $J = 7.3$ Hz, 2.0 Hz, 1H), 7.37 (dd, $J = 7.2$ Hz, 1.8 Hz, 1H), 7.28-7.21 (m, 2H), 2.25 (s, 2H), 1.86 (s, 3H).

^{13}C NMR (151 MHz, CDCl_3) δ 137.06, 132.47, 131.60, 129.98, 127.39, 126.63, 123.11, 51.52, 27.88.

HRMS (ESI): m/z Calcd. for $[\text{C}_9\text{H}_9\text{ClN}_2, \text{M}+\text{H}]^+$: 181.0527; Found: 181.0530.

2-amino-2-(2-methoxyphenyl)propanenitrile (**2h**)



The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.3$ in hexane: ethyl acetate = 2:1)

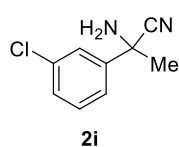
resulting in 42.9 mg of Colorless oil product **2h** with 61% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.39 (d, $J = 7.6$ Hz, 1H), 7.27 (dd, $J = 11.2$ Hz, 4.3 Hz, 1H), 6.91 (t, $J = 8.2$ Hz, 2H), 3.89 (s, 3H), 2.22 (s, 2H), 1.80 (s, 3H).

^{13}C NMR (151 MHz, CDCl_3) δ 157.11, 130.02, 128.60, 125.37, 124.22, 120.94, 112.00, 55.73, 49.91, 27.51.

HRMS (ESI): m/z Calcd. for $[\text{C}_{10}\text{H}_{12}\text{N}_2\text{O}, \text{M}+\text{H}]^+$: 177.1022; Found: 177.1018.

2-amino-2-(3-chlorophenyl)propanenitrile (**2i**)



The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.4$ in hexane:ethyl acetate

= 2:1) resulting in 51.1 mg of Colorless oil product **2i** with 71% isolated yield.

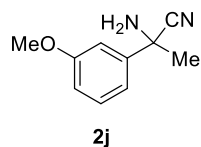
^1H NMR (600 MHz, CDCl_3) δ 7.60 (s, 1H), 7.48 (m, 1H), 7.30-7.24 (m, 2H), 2.06 (s, 2H), 1.68 (s, 3H).

^{13}C NMR (151 MHz, CDCl_3) δ 143.32, 134.90, 130.13, 128.89, 125.33, 123.60, 123.34,

53.29, 31.78.

HRMS (ESI): m/z Calcd. for $[C_9H_9ClN_2, M+H]^+$: 181.0527; Found: 181.0523.

2-amino-2-(3-methoxyphenyl)propanenitrile (**2j**)



The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) (R_f = 0.3 in hexane:ethyl acetate

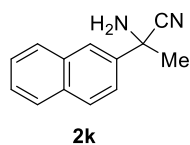
= 2:1) resulting in 48.6 mg of Colorless oil product **2j** with 69% isolated yield.

1H NMR (600 MHz, $CDCl_3$) δ 7.24 (t, J = 7.9 Hz, 1H), 7.17-7.09 (m, 2H), 6.81 (d, J = 8.0 Hz, 1H), 3.75 (s, 3H), 2.04 (s, 2H), 1.68 (s, 3H).

^{13}C NMR (151 MHz, $CDCl_3$) δ 159.99, 142.90, 129.91, 124.14, 117.25, 114.05, 110.77, 55.36, 53.57, 31.75.

HRMS (ESI): m/z Calcd. for $[C_{10}H_{12}N_2O, M+H]^+$: 177.1022; Found: 177.1017.

2-amino-2-(naphthalen-2-yl)propanenitrile (**2k**)



The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) (R_f = 0.3 in hexane:ethyl acetate

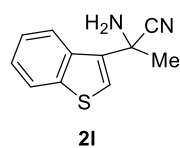
= 2:1) resulting in 42.3 mg of yellow solid product **2k** with 54% isolated yield.

1H NMR (600 MHz, $CDCl_3$) δ 8.06 (s, 1H), 7.80 (d, J = 8.5 Hz, 2H), 7.77-7.74 (m, 1H), 7.63 (dd, J = 8.7 Hz, 1.7 Hz, 1H), 7.46-7.41 (m, 2H), 2.13 (s, 2H), 1.76 (s, 3H).

^{13}C NMR (151 MHz, $CDCl_3$) δ 138.43, 133.23, 133.02, 128.91, 128.31, 127.64, 126.70, 126.69, 124.17, 122.69, 53.81, 31.63.

HRMS (ESI): m/z Calcd. for $[C_{13}H_{12}N_2, M+H]^+$: 197.1073; Found: 197.1067.

2-amino-2-(benzo[b]thiophen-3-yl)propanenitrile (**2l**)



The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) (R_f = 0.3 in hexane:ethyl acetate

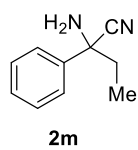
= 2:1) resulting in 41.2 mg of yellow solid product **2l** with 51% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.80 (d, $J = 7.3$ Hz, 1H), 7.74 (d, $J = 7.4$ Hz, 1H), 7.50 (s, 1H), 7.41-7.30 (m, 2H), 2.32 (s, 2H), 1.92 (s, 3H).

^{13}C NMR (151 MHz, CDCl_3) δ 146.47, 139.38, 138.93, 125.02, 124.75, 123.98, 122.77, 122.47, 121.39, 51.34, 32.10.

HRMS (ESI): m/z Calcd. for $[\text{C}_{11}\text{H}_{10}\text{N}_2\text{S}, \text{M}+\text{H}]^+$: 203.0637; Found: 203.0639.

2-amino-2-phenylbutanenitrile (**2m**)^[1]

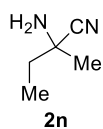


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.3$ in hexane:ethyl acetate = 2:1) resulting in 43.5 mg of colorless oil product **2m** with 68% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.59-7.52 (m, 2H), 7.33 (t, $J = 7.4$ Hz, 2H), 7.28 (t, $J = 7.3$ Hz, 1H), 2.04 (s, 2H), 1.95-1.83 (m, 2H), 0.91 (t, $J = 7.4$ Hz, 3H).

^{13}C NMR (151 MHz, CDCl_3) δ 139.95, 128.68, 128.61, 125.60, 123.27, 58.79, 37.25, 9.02.

2-amino-2-methylbutanenitrile (**2n**)^[5]

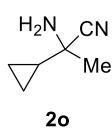


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.3$ in hexane:ethyl acetate = 2:1) resulting in 18.0 mg of colorless oil product **2n** with 46% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 1.73 (s, 2H), 1.69 (dd, $J = 14.0$ Hz, 6.8 Hz, 2H), 1.47 (s, 3H), 1.10 (t, $J = 7.5$ Hz, 3H).

^{13}C NMR (151 MHz, CDCl_3) δ 124.32, 50.51, 34.86, 27.14, 8.86.

2-amino-2-cyclopropylpropanenitrile (**2o**)^[6]



The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.3$ in hexane: ethyl acetate = 2:1) resulting in 18.9 mg of colorless oil product **2o** with 43% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 1.91 (s, 2H), 1.50 (s, 3H), 1.02 (m, 1H), 0.56-0.44 (m, 4H).

^{13}C NMR (151 MHz, CDCl_3) δ 120.62, 50.57, 26.01, 18.70, 8.75.

2-amino-3-chloro-2-methylpropanenitrile (**2p**)



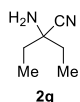
The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) (R_f = 0.3 in hexane:ethyl acetate = 2:1) resulting in 22.2 mg of colorless oil product **2p** with 47% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 3.65 (d, J = 11.1 Hz, 1H), 3.51 (d, J = 11.1 Hz, 1H), 2.09 (s, 2H), 1.59 (s, 3H).

^{13}C NMR (151 MHz, CDCl_3) δ 121.89, 51.47, 51.42, 25.70.

HRMS (ESI): m/z Calcd. for $[\text{C}_4\text{H}_7\text{ClN}_2, \text{M}+\text{H}]^+$: 119.0371; Found: 119.0371.

2-amino-2-ethylbutanenitrile (**2q**)^[3]

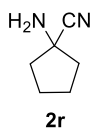


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) (R_f = 0.3 in hexane: ethyl acetate = 2:1) resulting in 25.5 mg of colorless oil product **2q** with 57% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 1.86 (s, 2H), 1.70 (dt, J = 14.8 Hz, 7.5 Hz, 2H), 1.57 (dt, J = 14.6 Hz, 7.4 Hz, 2H), 1.03 (t, J = 7.5 Hz, 6H).

^{13}C NMR (151 MHz, CDCl_3) δ 123.55, 54.98, 32.75, 8.44.

1-aminocyclopentane-1-carbonitrile (**2r**)^[7]

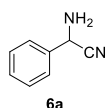


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) (R_f = 0.3 in hexane:ethyl acetate = 2:1) resulting in 27.7 mg of colorless oil product **2r** with 63% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 2.06 (s, 2H), 1.81 (d, J = 10.8 Hz, 2H), 1.79-1.73 (m, 4H), 1.70 (m, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 125.12, 54.57, 40.87, 23.63.

2-amino-2-phenylacetonitrile (**6a**)^[8]

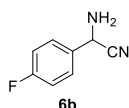


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) (R_f = 0.3 in hexane:ethyl acetate = 2:1) resulting in 47.0 mg of white solid product **6a** with 89% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.45 (d, J = 7.4 Hz, 2H), 7.32 (dt, J = 23.6 Hz, 7.2 Hz, 3H), 4.82 (s, 1H), 1.89 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 136.36, 129.12, 129.05, 126.66, 120.92, 47.32.

2-amino-2-(4-fluorophenyl)acetonitrile (**6b**)^[9]

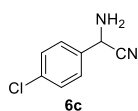


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) (R_f = 0.3 in hexane:ethyl acetate = 2:1) resulting in 53.4 mg of yellow solid product **6b** with 89% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.55 (dd, J = 7.3, 5.3 Hz, 2H), 7.13 (t, J = 8.3 Hz, 2H), 4.93 (s, 1H), 1.96 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 163.82, 162.17, 128.54 (d, J = 8.4 Hz), 120.72, 116.05 (d, J = 21.9 Hz), 46.64.

2-amino-2-(4-chlorophenyl)acetonitrile (**6c**)^[10]

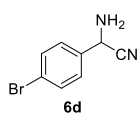


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) (R_f = 0.3 in hexane:ethyl acetate = 2:1) resulting in 57.8 mg of yellow solid product **6c** with 87% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.41 (d, J = 8.4 Hz, 2H), 7.32 (d, J = 8.4 Hz, 2H), 4.82 (s, 1H), 1.88 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 135.07, 134.79, 129.26, 128.07, 120.54, 46.69.

2-amino-2-(4-bromophenyl)acetonitrile (**6d**)^[10]

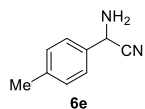


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.3$ in hexane:ethyl acetate = 2:1) resulting in 69.7 mg of yellow solid product **6d** with 83% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.48 (d, $J = 8.0$ Hz, 2H), 7.35 (d, $J = 7.8$ Hz, 2H), 4.80 (s, 1H), 1.85 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 135.30, 132.23, 128.36, 123.19, 120.45, 46.75.

2-amino-2-(p-tolyl)acetonitrile (**6e**)^[10]

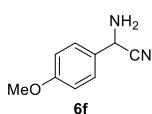


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.3$ in hexane:ethyl acetate = 2:1) resulting in 53.1 mg of yellow solid product **6e** with 91% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.32 (d, $J = 7.8$ Hz, 2H), 7.13 (d, $J = 7.7$ Hz, 2H), 4.77 (s, 1H), 2.28 (s, 3H), 1.84 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 138.95, 133.57, 129.74, 126.55, 121.09, 47.08, 21.11.

2-amino-2-(4-methoxyphenyl)acetonitrile (**6f**)^[10]

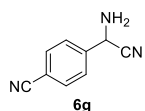


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.2$ in hexane:ethyl acetate = 2:1) resulting in 24.6 mg of yellow solid product **6f** with 38% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.37 (d, $J = 8.6$ Hz, 2H), 6.86 (d, $J = 8.7$ Hz, 2H), 4.79 (s, 1H), 3.75 (s, 3H), 1.79 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 160.13, 128.52, 127.94, 121.07, 114.46, 55.38, 46.77.

4-(amino(cyano)methyl)benzonitrile (**6g**)^[11]

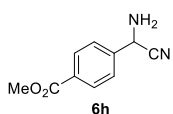


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.2$ in hexane:ethyl acetate = 2:1) resulting in 36.4 mg of yellow solid product **6g** with 58% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.65 (q, $J = 8.5$ Hz, 4H), 4.92 (s, 1H), 1.94 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 141.13, 132.83, 127.55, 119.91, 118.06, 113.16, 46.85.

methyl 4-(amino(cyano)methyl)benzoate (**6h**)^[11]

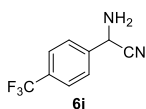


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.2$ in hexane:ethyl acetate = 2:1) resulting in 40.3 mg of yellow solid product **6h** with 53% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 8.01 (d, $J = 8.3$ Hz, 2H), 7.56 (d, $J = 8.2$ Hz, 2H), 4.90 (s, 1H), 3.86 (s, 3H), 1.94 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 166.35, 140.89, 130.90, 130.32, 126.73, 120.38, 52.29, 47.03.

2-amino-2-(4-(trifluoromethyl)phenyl)acetonitrile (**6i**)^[12]

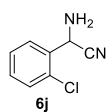


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.2$ in hexane:ethyl acetate = 2:1) resulting in 64.8 mg of yellow solid product **6i** with 81% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.51-7.36 (m, 2H), 7.03 (t, $J = 8.4$ Hz, 2H), 4.82 (s, 1H), 1.83 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 163.81, 162.17, 132.18, 128.55 (q, $J = 8.4$ Hz), 120.74, 116.04 (q, $J = 21.9$ Hz), 46.63.

2-amino-2-(2-chlorophenyl)acetonitrile (**6j**)^[10]

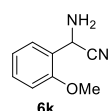


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.3$ in hexane:ethyl acetate = 2:1) resulting in 44.5 mg of yellow solid product **6j** with 67% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.68-7.63 (m, 1H), 7.44 (d, $J = 7.1$ Hz, 1H), 7.39-7.32 (m, 2H), 5.25 (s, 1H), 2.04 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 134.45, 132.82, 130.48, 130.32, 128.17, 127.78, 119.83, 45.19.

2-amino-2-(2-methoxyphenyl)acetonitrile (**6k**)^[13]

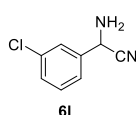


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.2$ in hexane:ethyl acetate = 2:1) resulting in 48.0 mg of yellow solid product **6k** with 74% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.44 (dd, $J = 7.5$ Hz, 1.4 Hz, 1H), 7.38 (dt, $J = 8.2$ Hz, 1.6 Hz, 1H), 7.02 (dt, $J = 7.5$ Hz, 0.7 Hz, 1H), 6.96 (d, $J = 8.2$ Hz, 1H), 5.07 (s, 1H), 3.92 (s, 3H), 2.16 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 156.64, 130.43, 127.74, 125.27, 121.06, 120.74, 111.23, 55.65, 43.51.

2-amino-2-(3-chlorophenyl)acetonitrile (**6l**)^[11]

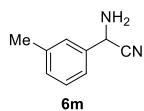


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.2$ in hexane:ethyl acetate = 2:1) resulting in 59.1 mg of yellow solid product **6l** with 89% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.49 (s, 1H), 7.38-7.34 (m, 1H), 7.29 (d, $J = 4.8$ Hz, 2H), 4.83 (s, 1H), 1.85 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 138.16, 135.09, 130.35, 129.28, 126.96, 124.82, 120.35, 46.76.

2-amino-2-(m-tolyl)acetonitrile (**6m**)^[10]

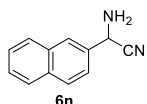


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.2$ in hexane:ethyl acetate = 2:1) resulting in 42.0 mg of yellow solid product **6m** with 72% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.32 (m, 3H), 7.19 (d, $J = 6.1$ Hz, 1H), 4.87 (s, 1H), 2.39 (s, 3H), 1.93 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 139.03, 136.30, 129.78, 129.01, 127.30, 123.68, 120.99, 47.30, 21.36.

2-amino-2-(naphthalen-2-yl)acetonitrile (**6n**)



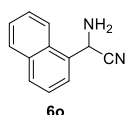
The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.4$ in hexane:ethyl acetate = 2:1) resulting in 51.0 mg of white solid product **6n** with 70% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.93 (s, 1H), 7.82-7.78 (m, 3H), 7.51 (d, $J = 8.2$ Hz, 1H), 7.45 (dd, $J = 5.6$ Hz, 2.9 Hz, 2H), 4.98 (s, 1H), 1.89 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 133.63, 133.37, 133.14, 129.17, 128.17, 127.75, 126.87, 126.80, 125.73, 124.17, 120.89, 47.49.

HRMS (ESI): m/z Calcd. for $[\text{C}_{12}\text{H}_{10}\text{N}_2, \text{M}+\text{H}]^+$: 183.0917; Found: 183.0912.

2-amino-2-(naphthalen-1-yl)acetonitrile (**6o**)^[14]



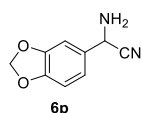
The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.4$ in hexane:ethyl acetate = 2:1) resulting in 52.4 mg of white solid product **6o** with 72% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 8.16 (d, $J = 8.5$ Hz, 1H), 7.93 (t, $J = 9.1$ Hz, 2H), 7.83 (d, $J = 7.1$ Hz, 1H), 7.64 (t, $J = 7.6$ Hz, 1H), 7.58 (t, $J = 7.5$ Hz, 1H), 7.53 (t, $J = 7.7$ Hz, 1H), 5.61 (s, 1H), 2.01 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 134.12, 131.74, 130.09, 130.03, 129.12, 127.12, 126.34,

125.26, 124.96, 122.82, 120.95, 45.39.

2-amino-2-(benzo[d][1,3]dioxol-5-yl)acetonitrile (**6p**)^[15]

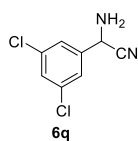


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.2$ in hexane:ethyl acetate = 2:1) resulting in 57.7 mg of yellow solid product **6p** with 82% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 6.99 (d, $J = 8.0$ Hz, 2H), 6.84-6.79 (m, 1H), 5.99 (d, $J = 0.9$ Hz, 2H), 4.81 (s, 1H), 1.89 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 148.31, 148.19, 130.28, 120.96, 120.23, 108.49, 107.23, 101.51, 47.02.

2-amino-2-(3,5-dichlorophenyl)acetonitrile (**6q**)



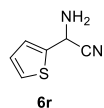
The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.2$ in hexane:ethyl acetate = 2:1) resulting in 72.8 mg of yellow solid product **6q** with 91% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.39 (d, $J = 1.0$ Hz, 2H), 7.31 (s, 1H), 4.81 (s, 1H), 1.92 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 139.38, 135.76, 129.28, 125.34, 119.86, 46.33.

HRMS (ESI): m/z Calcd. for $[\text{C}_8\text{H}_6\text{Cl}_2\text{N}_2, \text{M}+\text{H}]^+$: 200.9981; Found: 200.9975.

2-amino-2-(thiophen-2-yl)acetonitrile (**6r**)^[10]

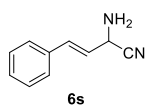


The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.3$ in hexane:ethyl acetate = 2:1) resulting in 50.2 mg of yellow solid product **6r** with 58% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.29-7.24 (m, 1H), 7.16 (dd, $J = 2.4$ Hz, 1.1 Hz, 1H), 6.93 (dd, $J = 5.0$ Hz, 3.7 Hz, 1H), 5.05 (s, 1H), 1.97 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 139.82, 127.01, 126.62, 125.94, 120.08, 43.38.

2-amino-4-phenylbut-3-enenitrile (**6s**)



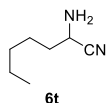
The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.3$ in hexane:ethyl acetate = 2:1) resulting in 34.1 mg of white solid product **6s** with 54% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 7.31 (t, $J = 7.5$ Hz, 2H), 7.24 (dd, $J = 12.4$ Hz, 5.0 Hz, 1H), 7.19 (d, $J = 7.5$ Hz, 2H), 5.42 (t, $J = 7.5$ Hz, 1H), 3.39 (d, $J = 7.5$ Hz, 2H), 3.28 (s, 2H).

^{13}C NMR (151 MHz, CDCl_3) δ 138.07, 128.84, 128.21, 126.75, 117.79, 117.47, 116.67, 43.45.

HRMS (ESI): m/z Calcd. for $[\text{C}_{10}\text{H}_{10}\text{N}_2, \text{M}+\text{H}]^+$: 159.0917; Found: 159.0920.

2-aminoheptanenitrile (**6t**)



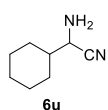
The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.3$ in hexane:ethyl acetate = 2:1) resulting in 43.8 mg of Colorless oil product **6t** with 87% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 3.68 (t, $J = 7.1$ Hz, 1H), 1.74 (dd, $J = 15.1$ Hz, 7.8 Hz, 2H), 1.63 (s, 2H), 1.55-1.46 (m, 2H), 1.34 (dd, $J = 6.8$ Hz, 3.3 Hz, 4H), 0.91 (t, $J = 6.8$ Hz, 3H).

^{13}C NMR (151 MHz, CDCl_3) δ 122.23, 43.44, 35.33, 31.13, 25.11, 22.38, 13.89.

HRMS (ESI): m/z Calcd. for $[\text{C}_7\text{H}_{14}\text{N}_2, \text{M}+\text{H}]^+$: 127.1230; Found: 127.1231.

2-amino-2-cyclohexylacetonitrile (**6u**)^[16]



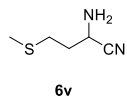
The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.3$ in hexane:ethyl acetate = 2:1) resulting in 46.9 mg of Colorless oil product **6u** with 85% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 3.52 (d, $J = 6.0$ Hz, 1H), 1.91-1.78 (m, 4H), 1.71 (m,

1H), 1.63 (s, 2H), 1.34 (d, $J = 3.4$ Hz, 1H), 1.29-1.25 (m, 2H), 1.22-1.10 (m, 3H).

^{13}C NMR (151 MHz, CDCl_3) δ 121.42, 49.02, 42.14, 29.28, 28.27, 25.98.

2-amino-4-(methylthio)butanenitrile (**6v**)^[17]



The title compound was prepared according to the general procedure as described, silica gel flash column chromatography was performed using hexanes and ethyl acetate (4:1) ($R_f = 0.3$ in hexane:ethyl acetate = 2:1)

resulting in 19.2 mg of yellow oil product **6v** with 37% isolated yield.

^1H NMR (600 MHz, CDCl_3) δ 3.93 (t, $J = 7.1$ Hz, 1H), 2.75-2.71 (m, 1H), 2.68-2.66 (m, 1H), 2.13 (s, 3H), 2.07-1.99 (m, 2H), 1.84 (s, 2H).

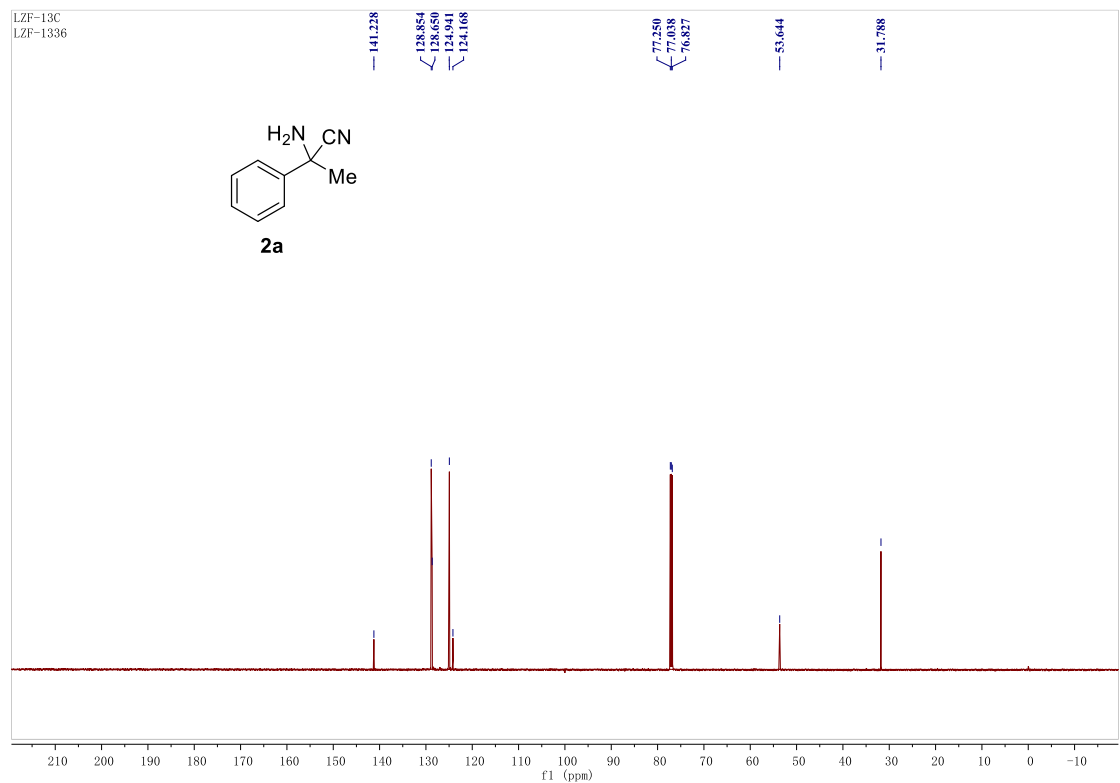
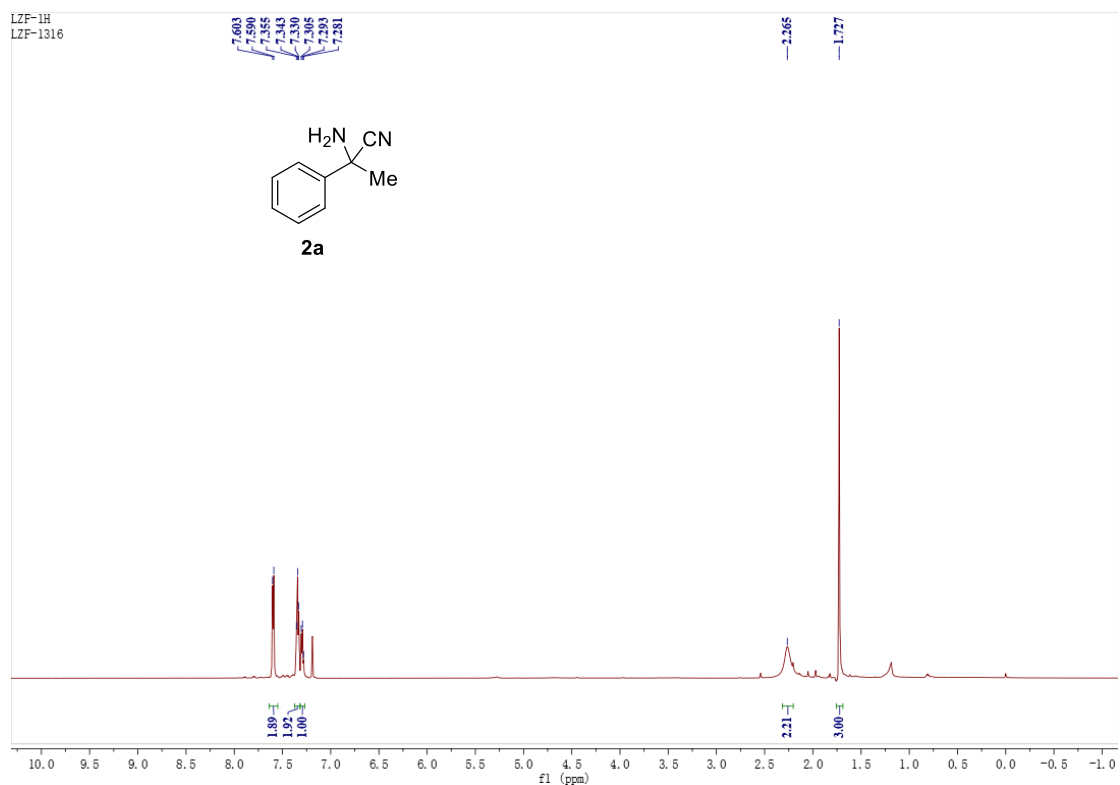
^{13}C NMR (151 MHz, CDCl_3) δ 121.79, 41.99, 34.42, 29.89, 15.45.

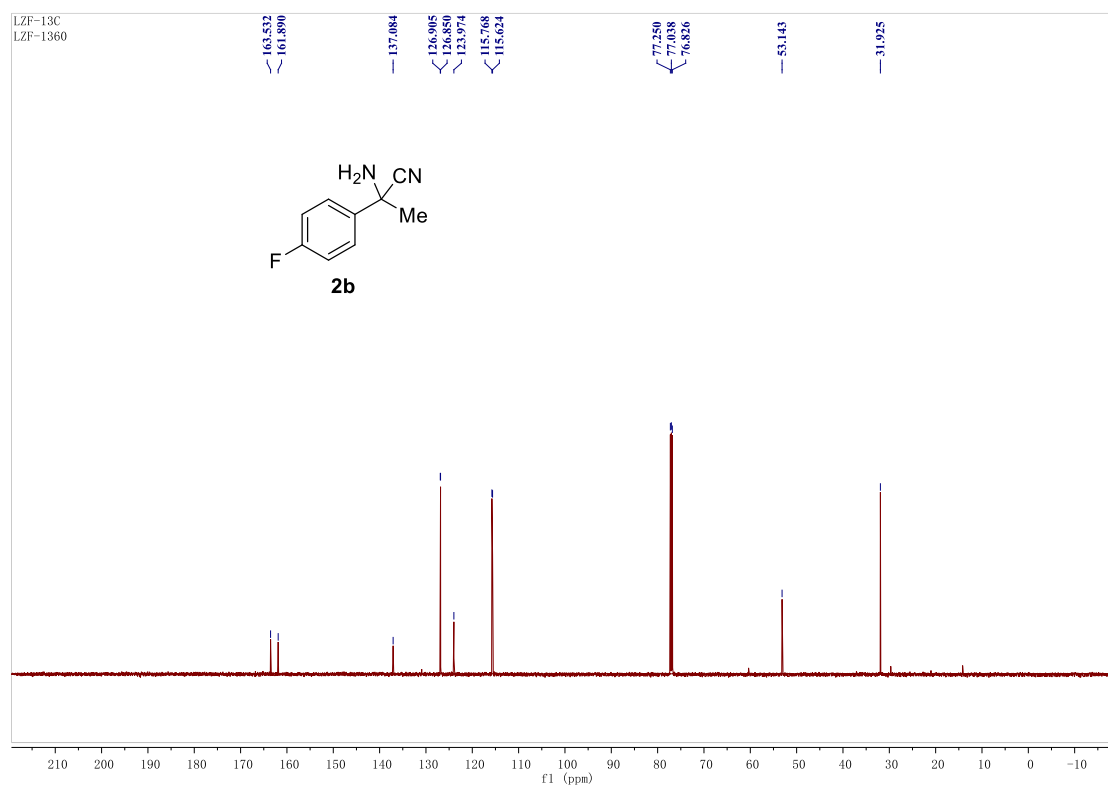
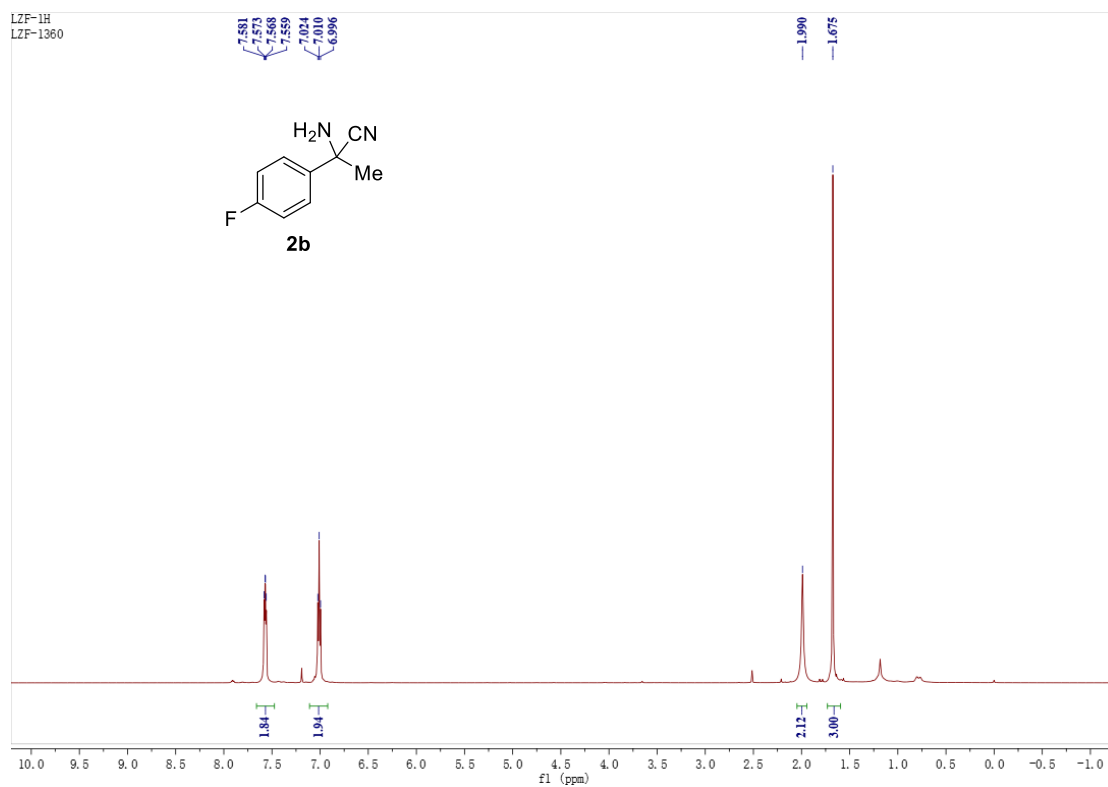
V References

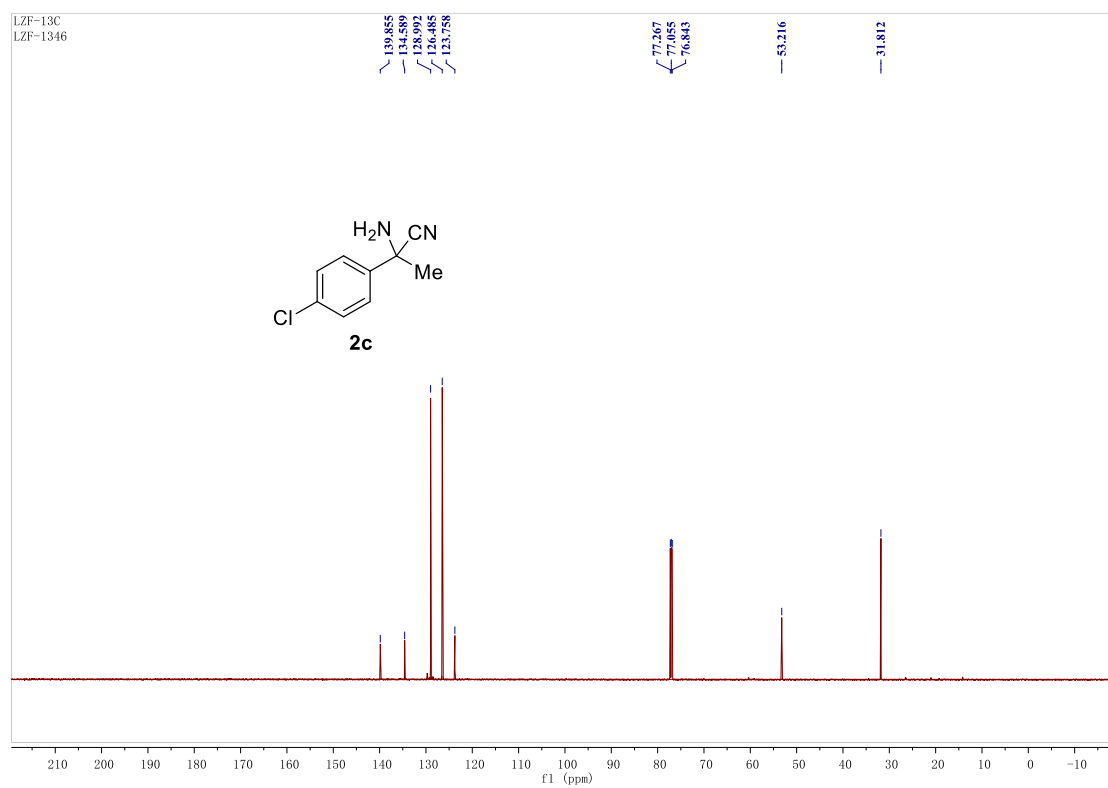
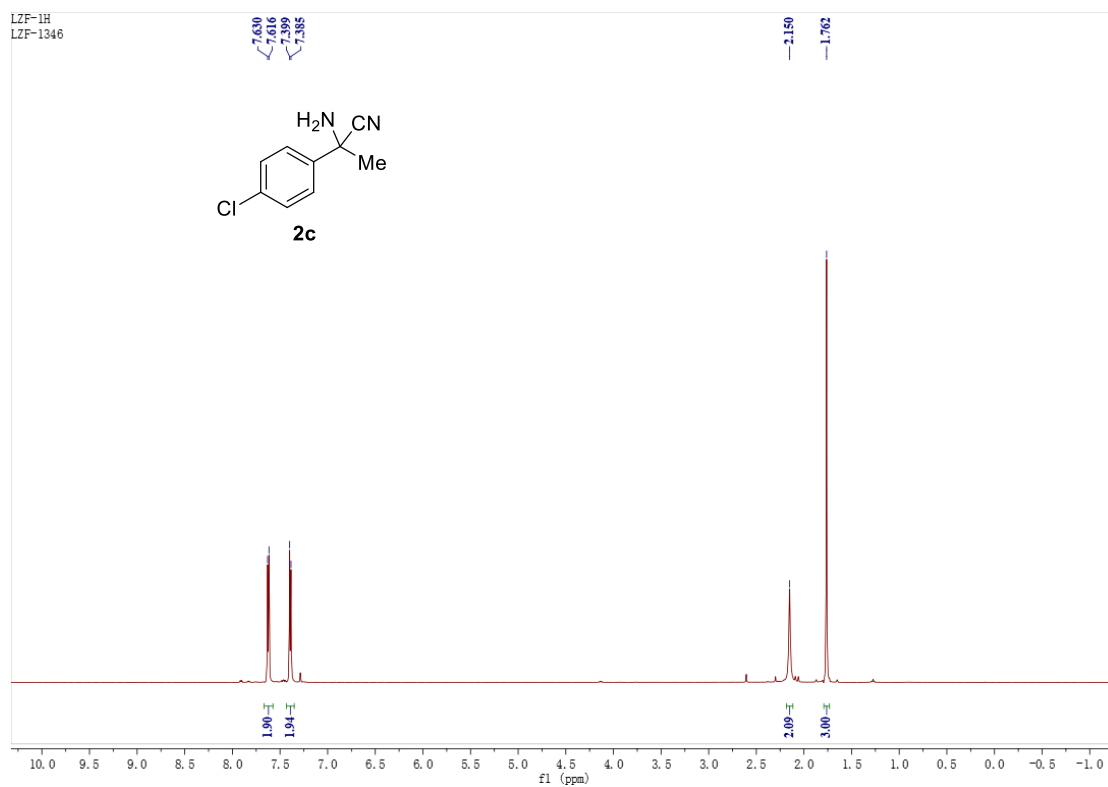
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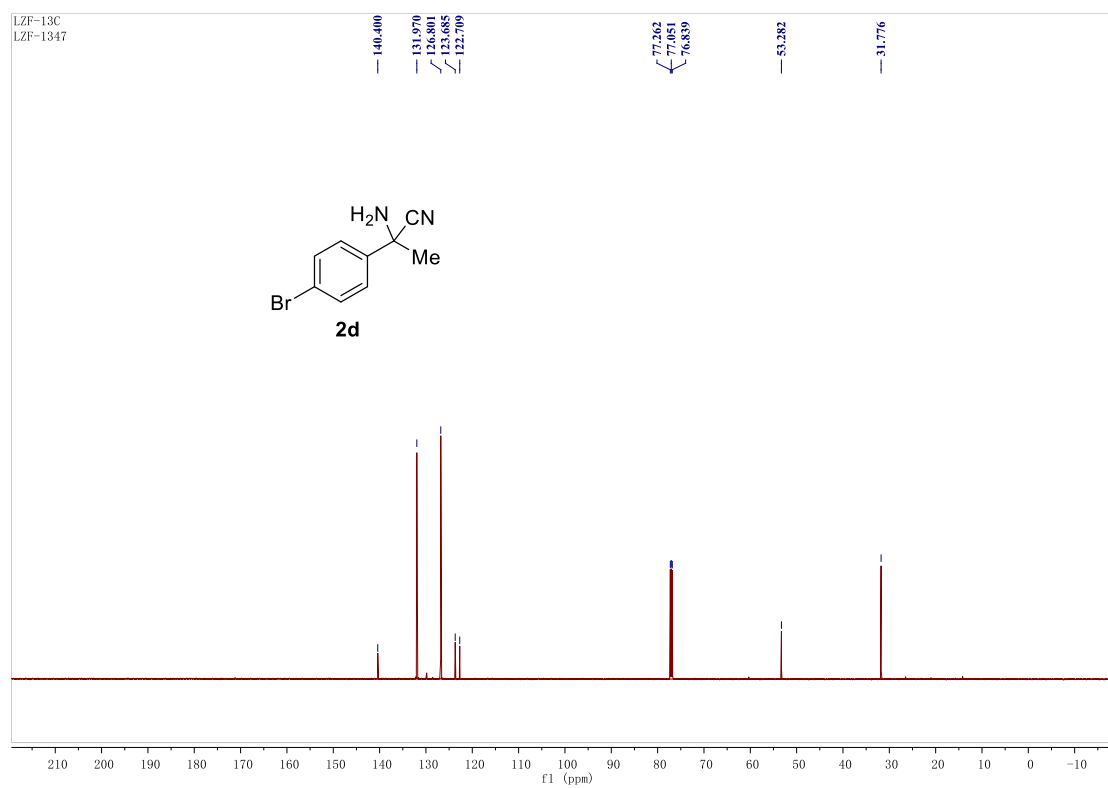
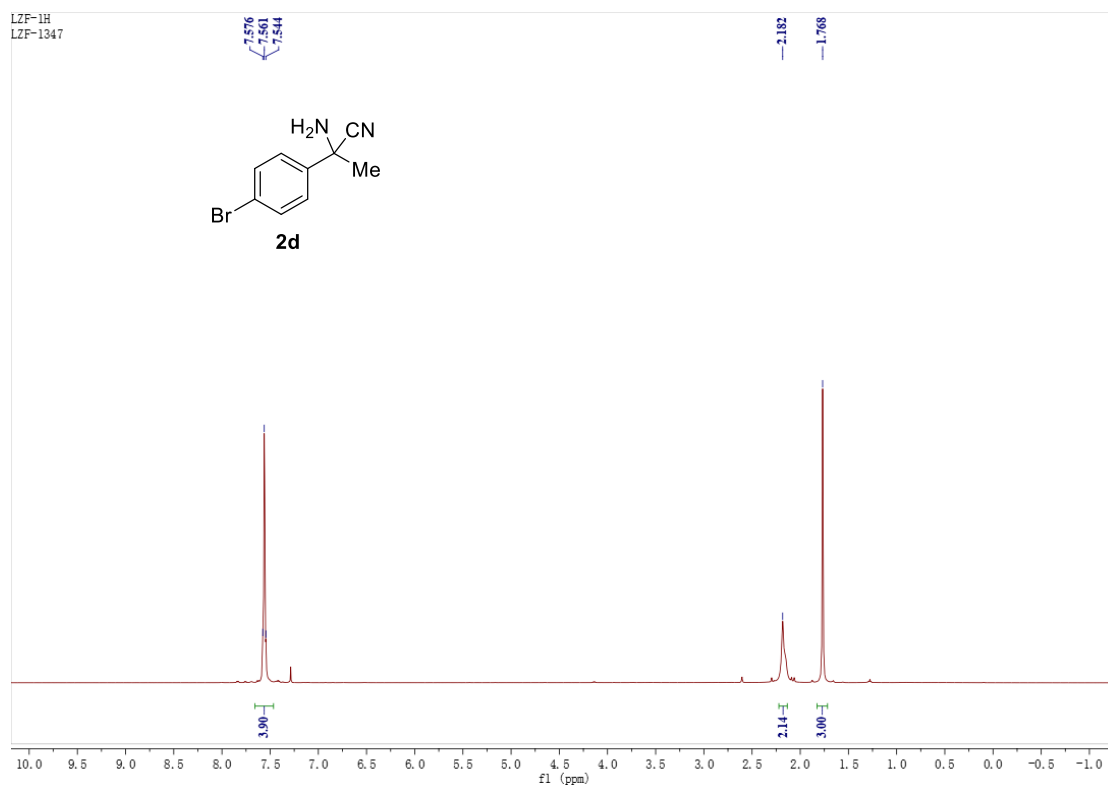
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VI NMR spectra of the products





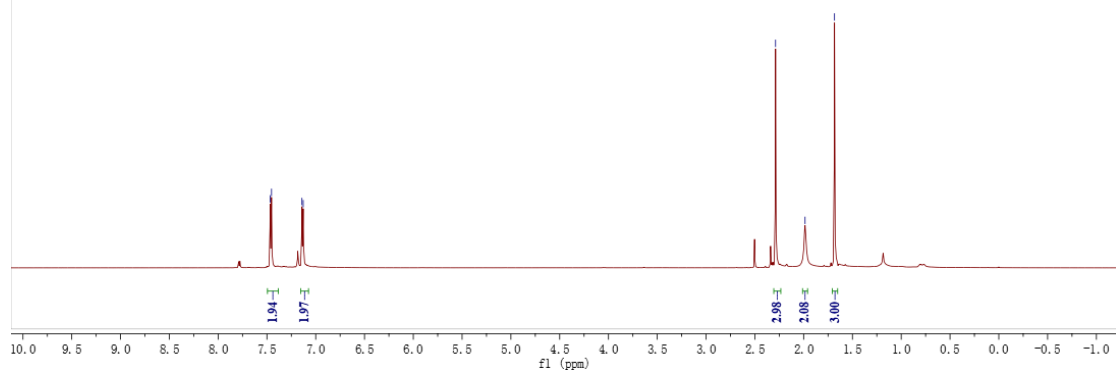
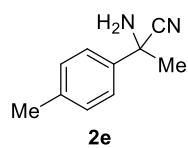




LZF-1H
LZF-1372

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7.453
7.141
7.128

2.238
1.985
1.684



LZF-13C
LZF-1372

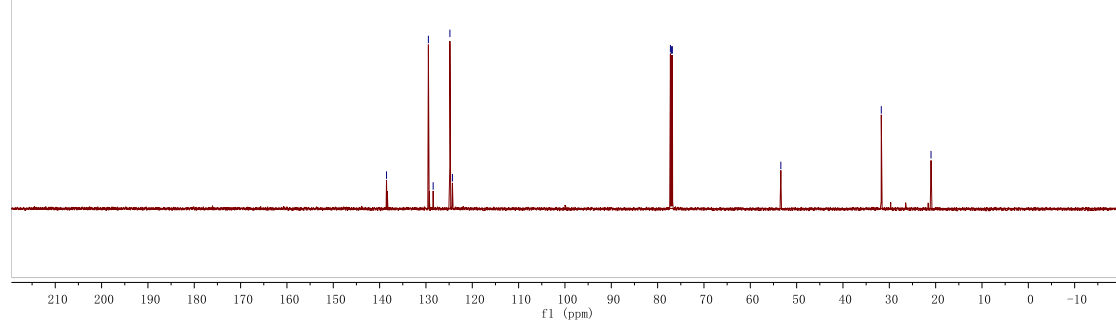
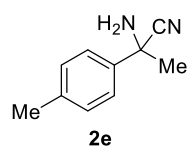
138.509
129.474
128.443
124.831
124.287

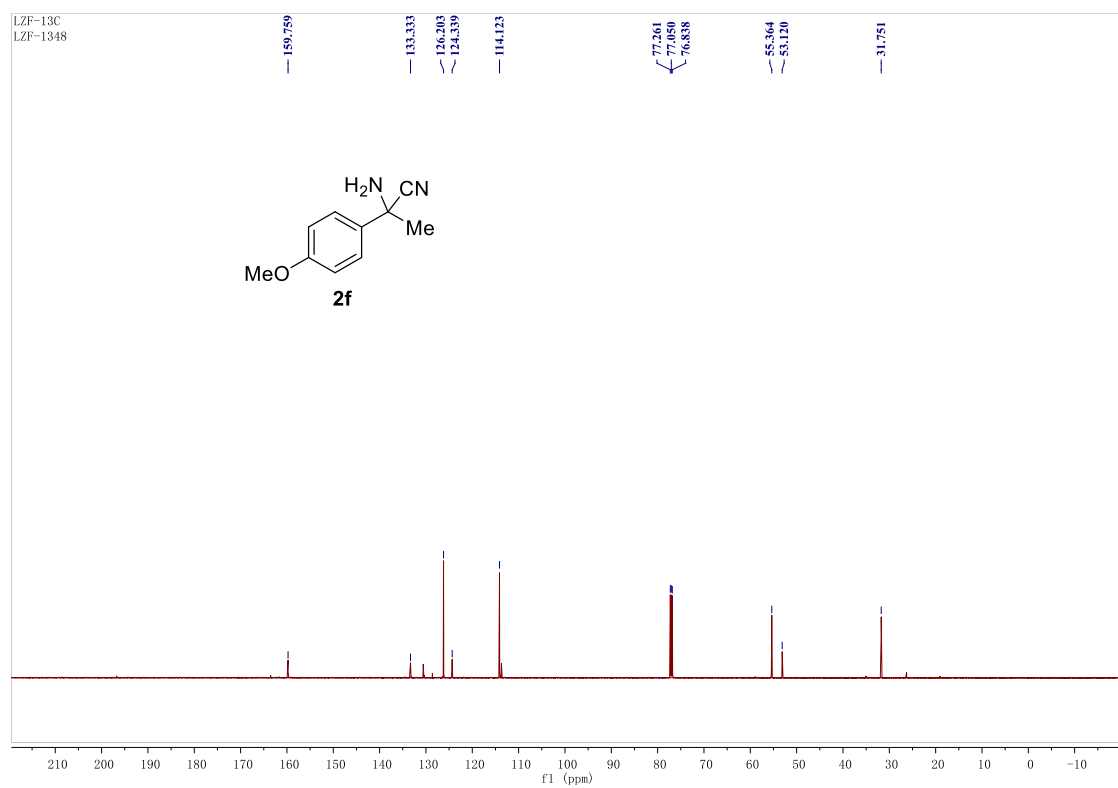
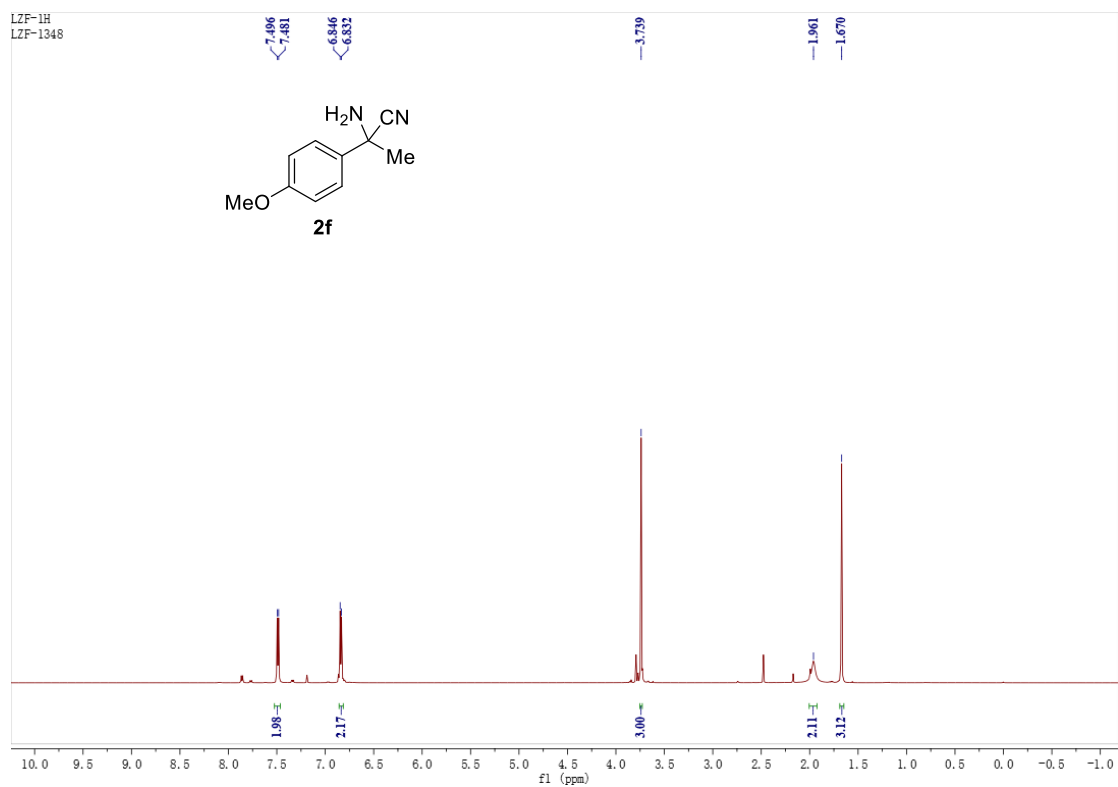
77.241
77.029
76.817

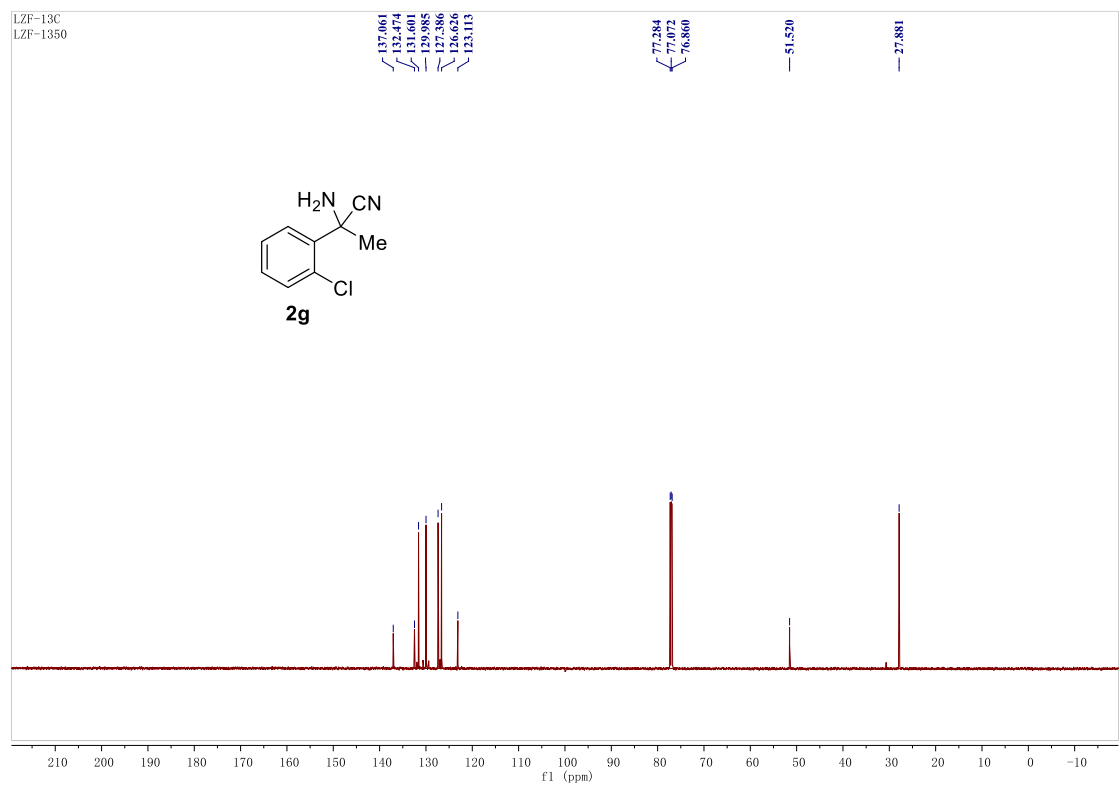
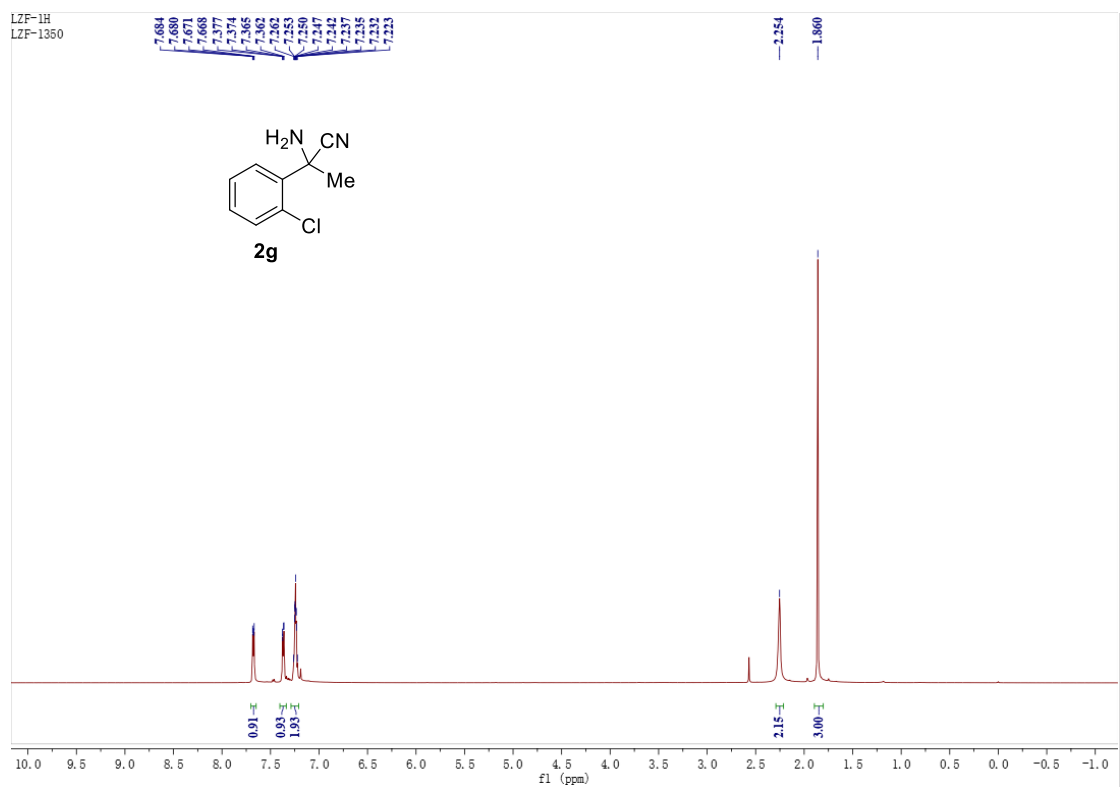
53.410

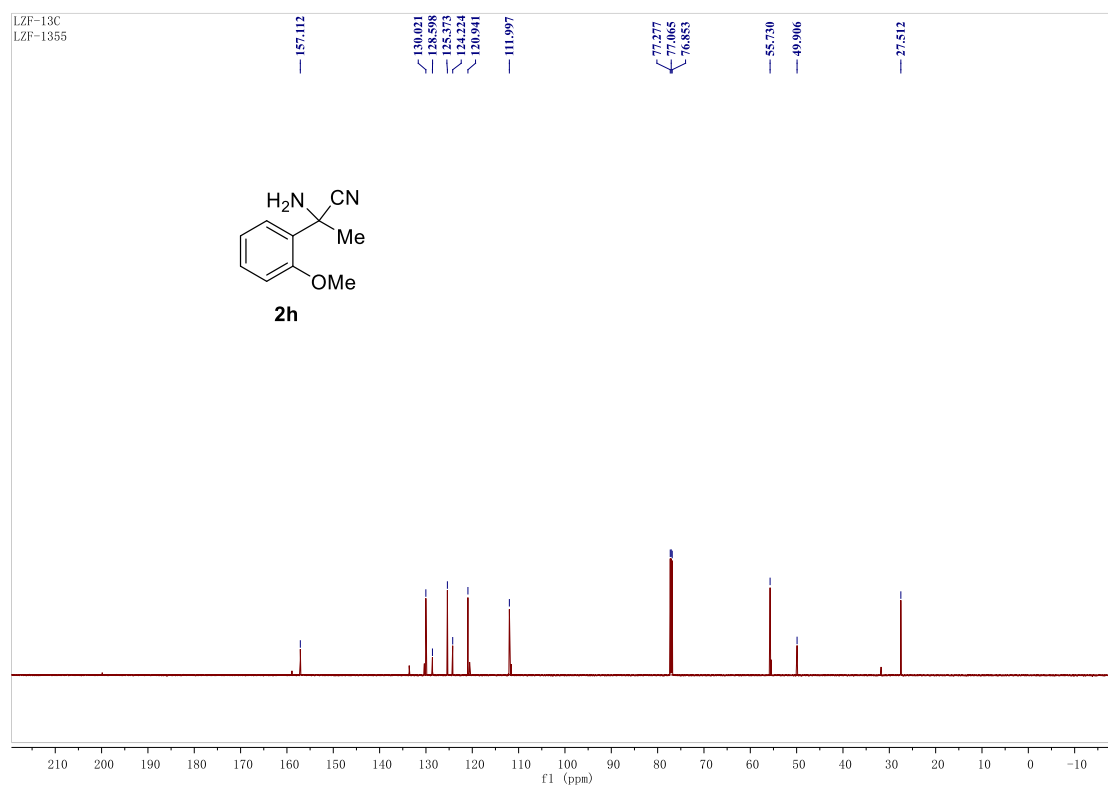
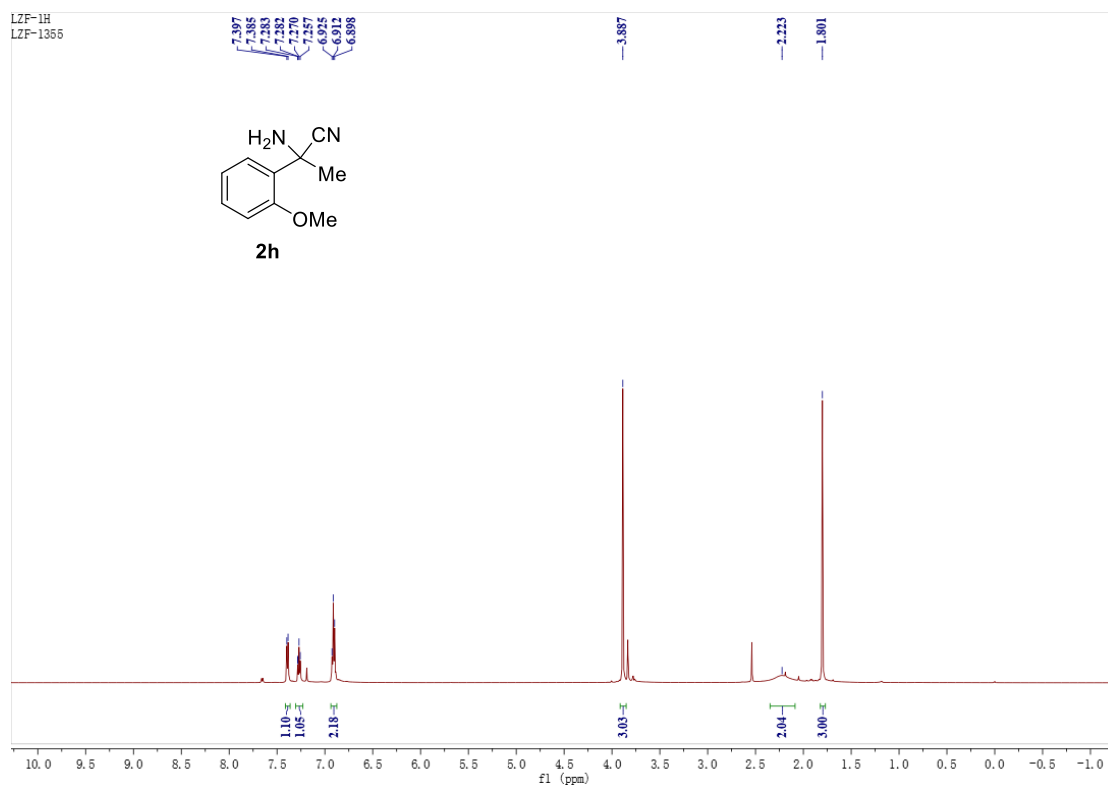
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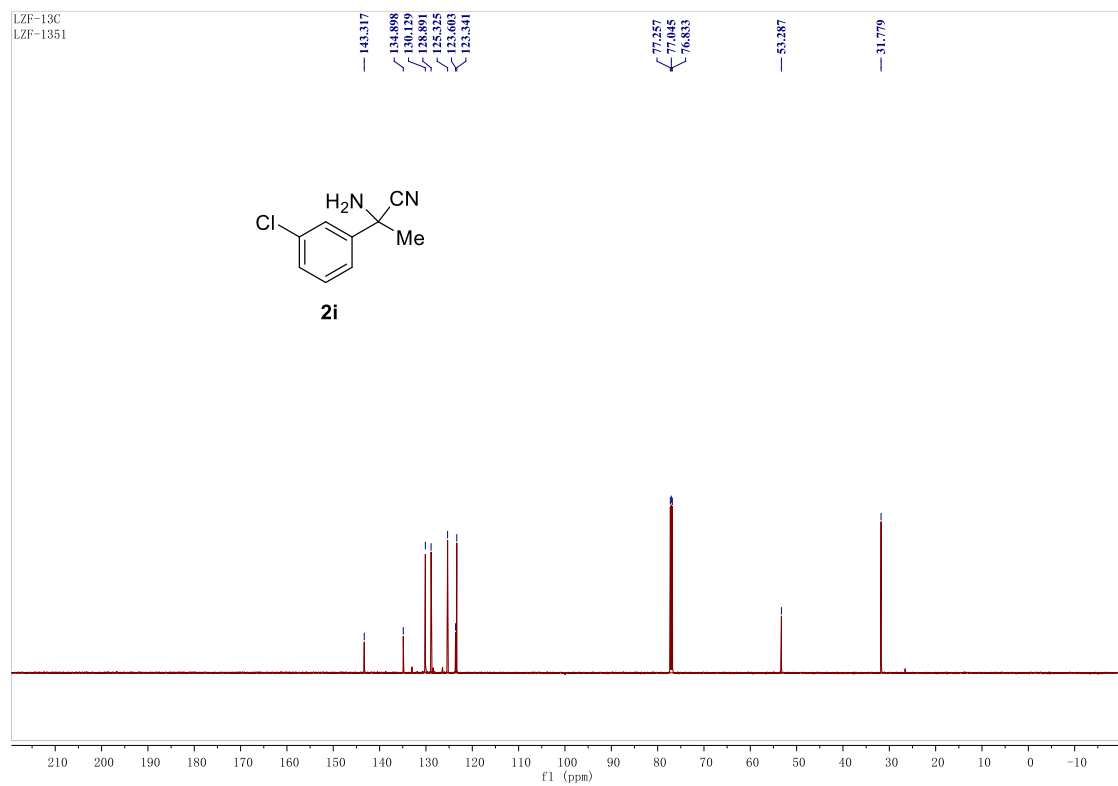
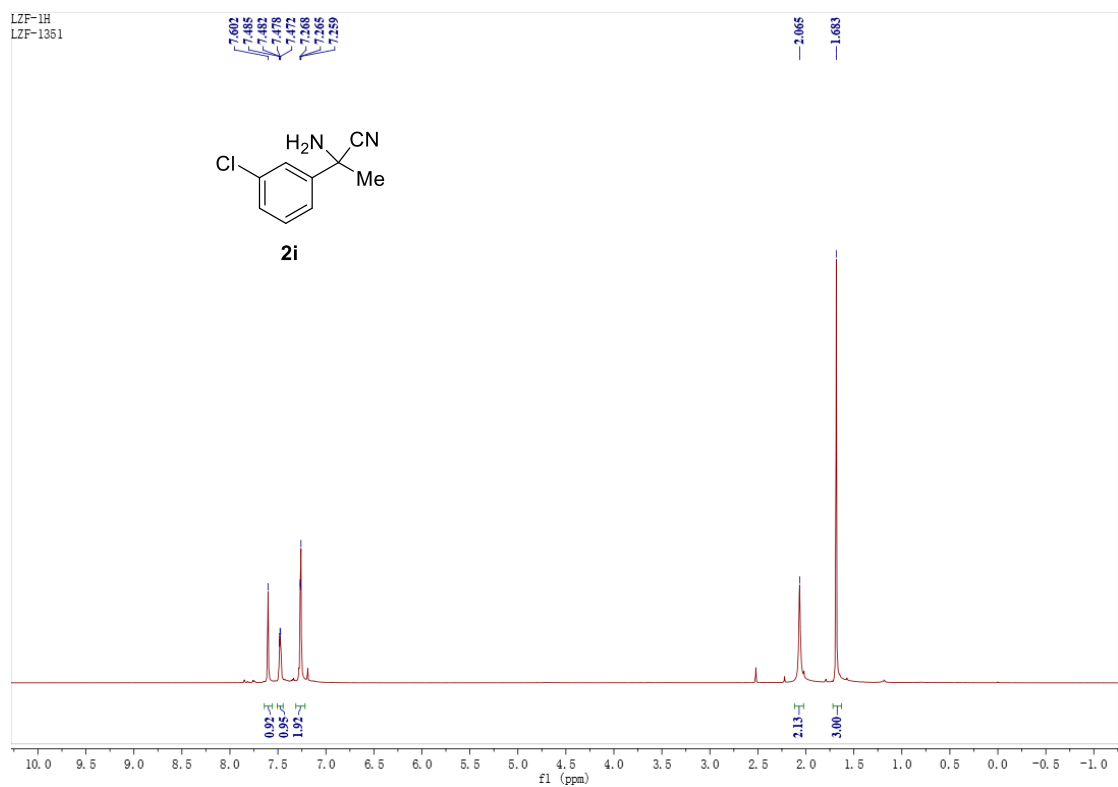
20.998

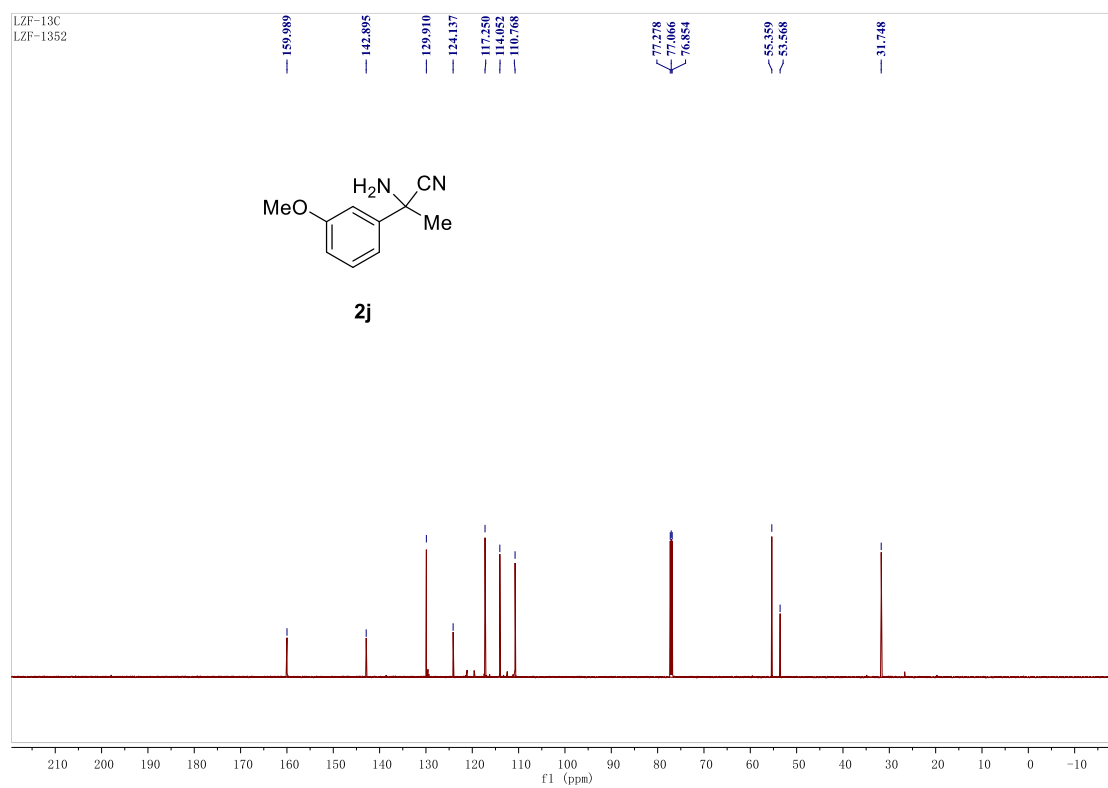
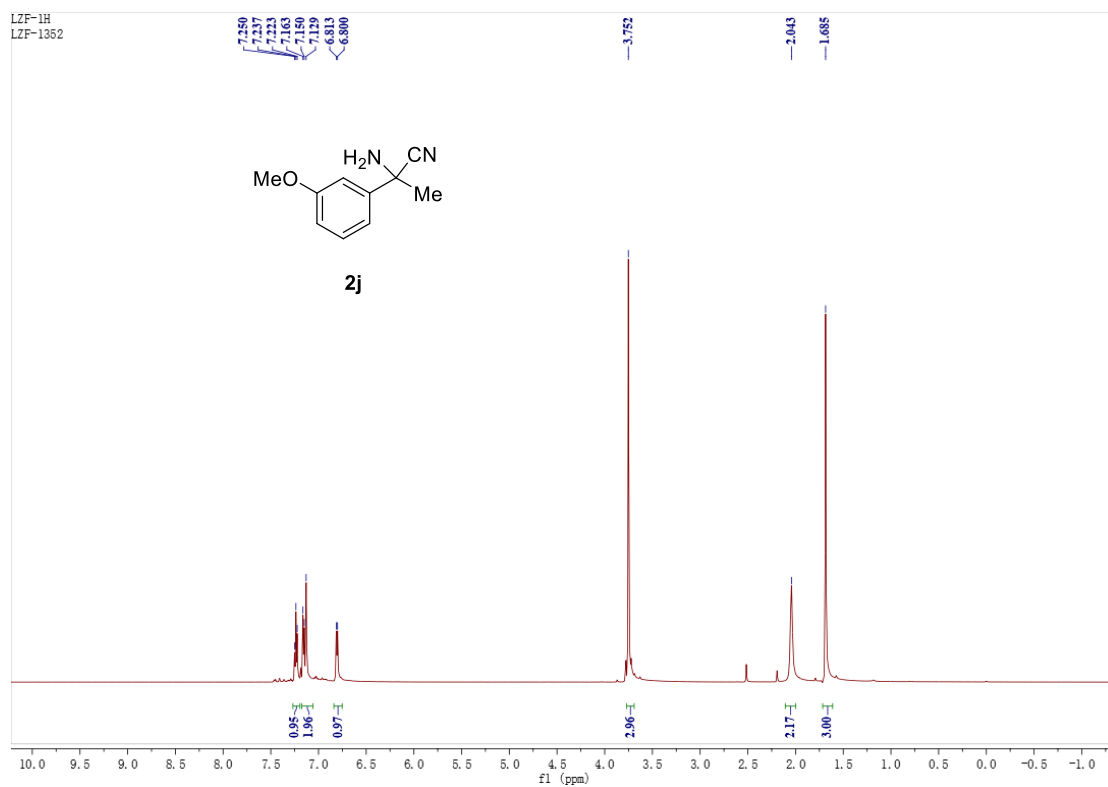


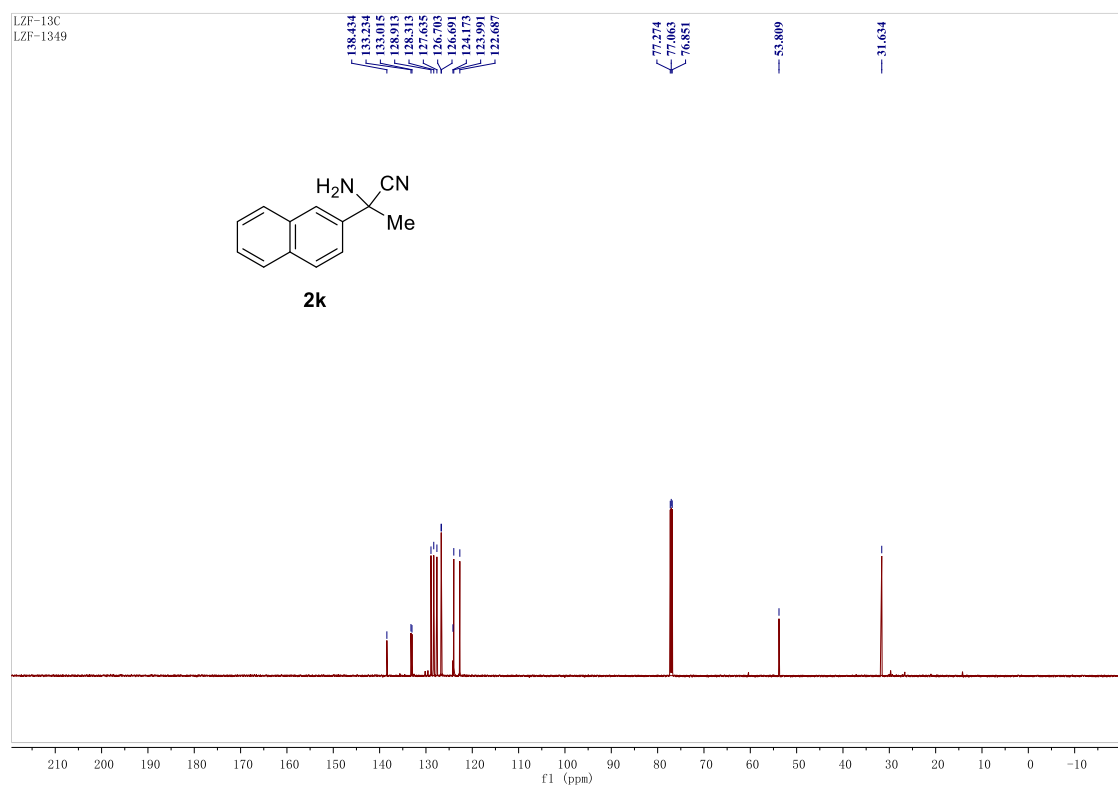
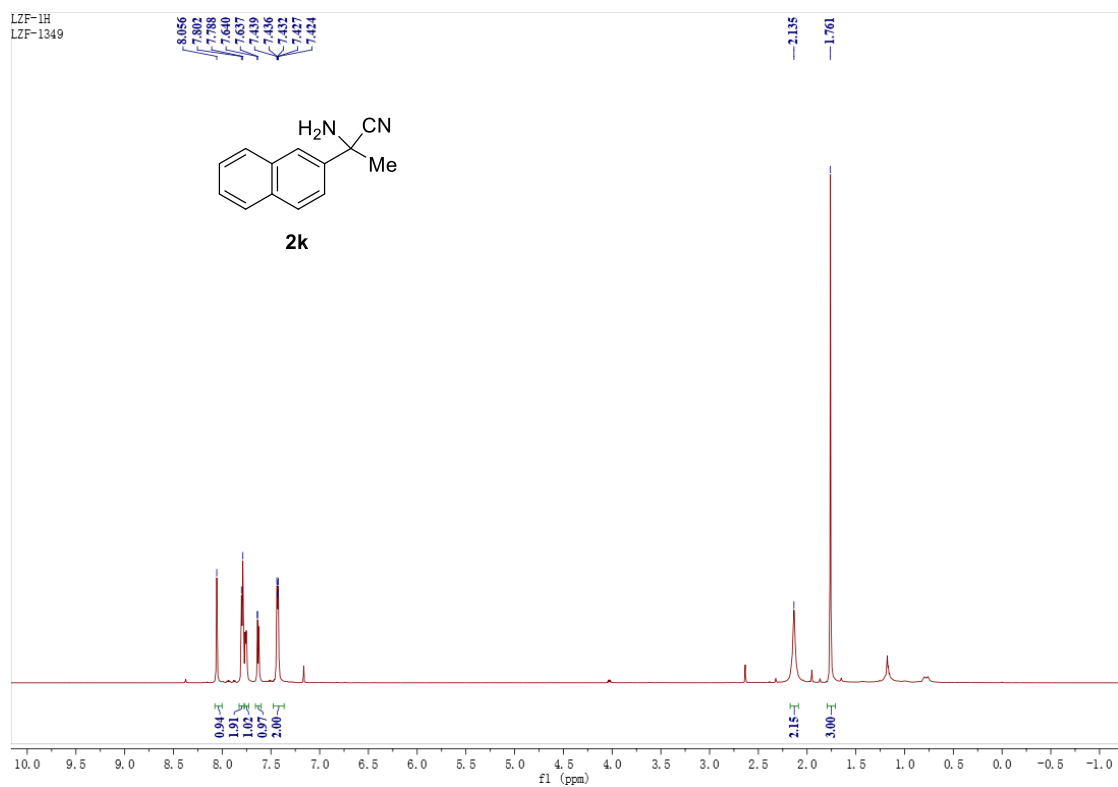


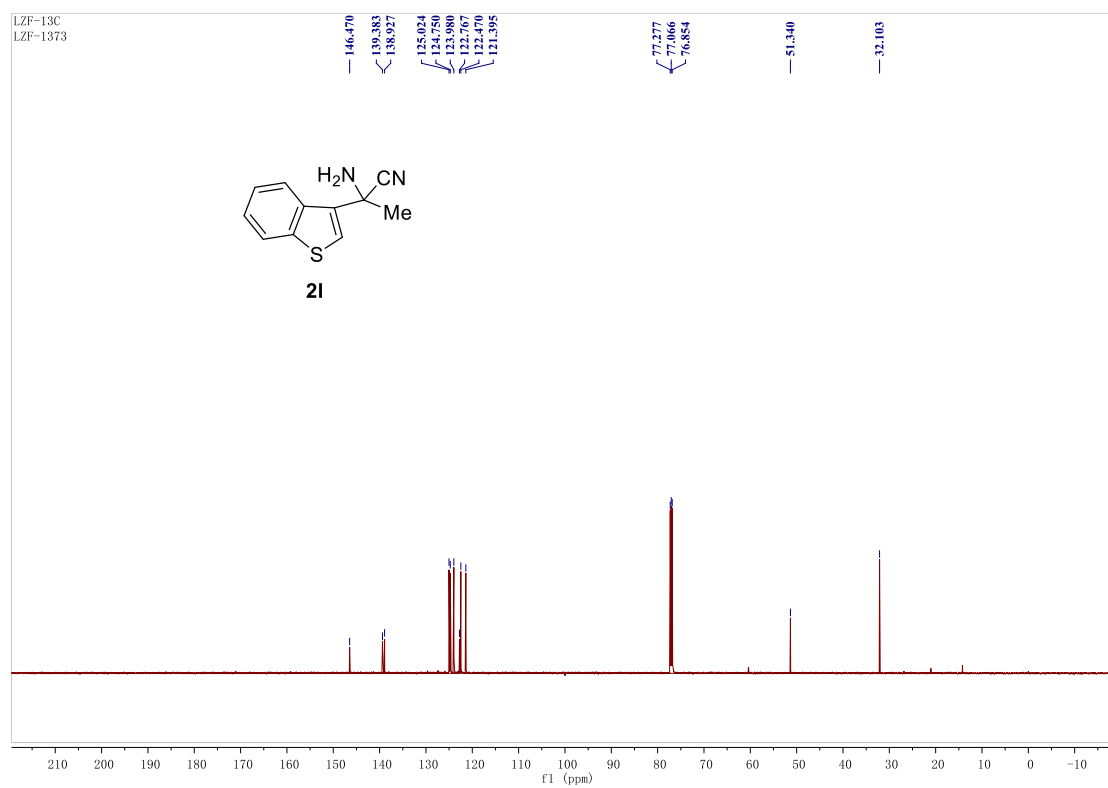
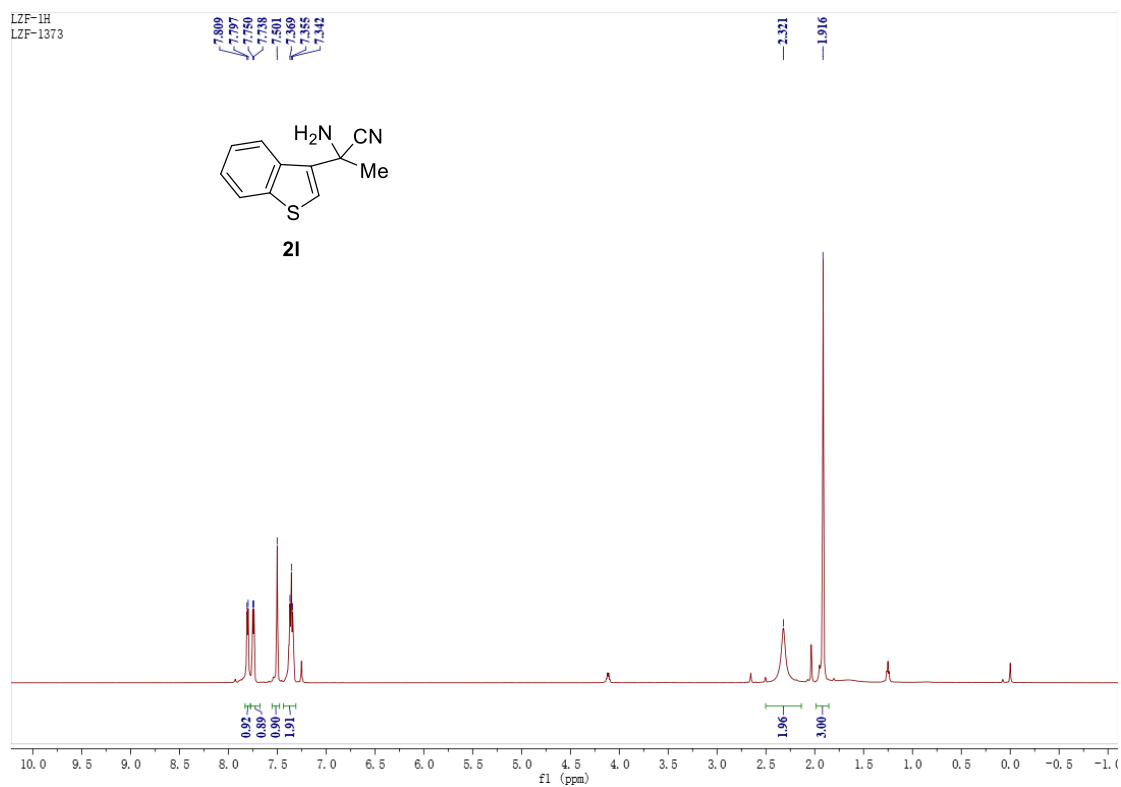


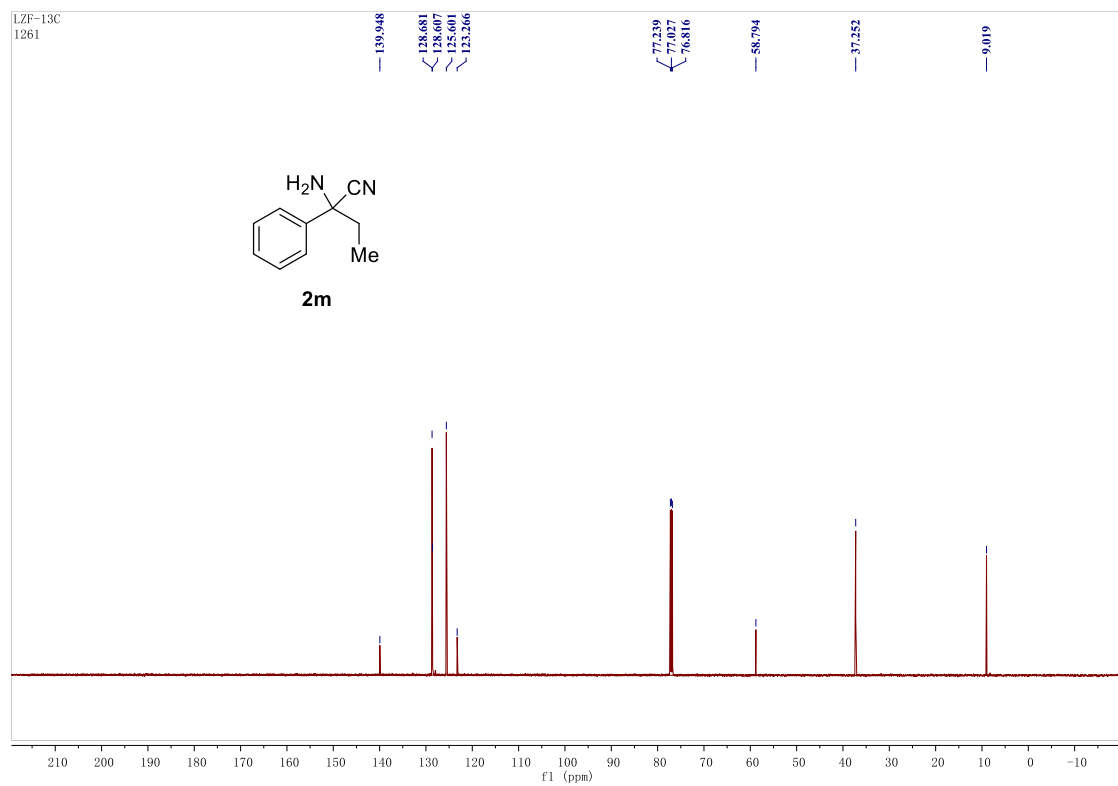
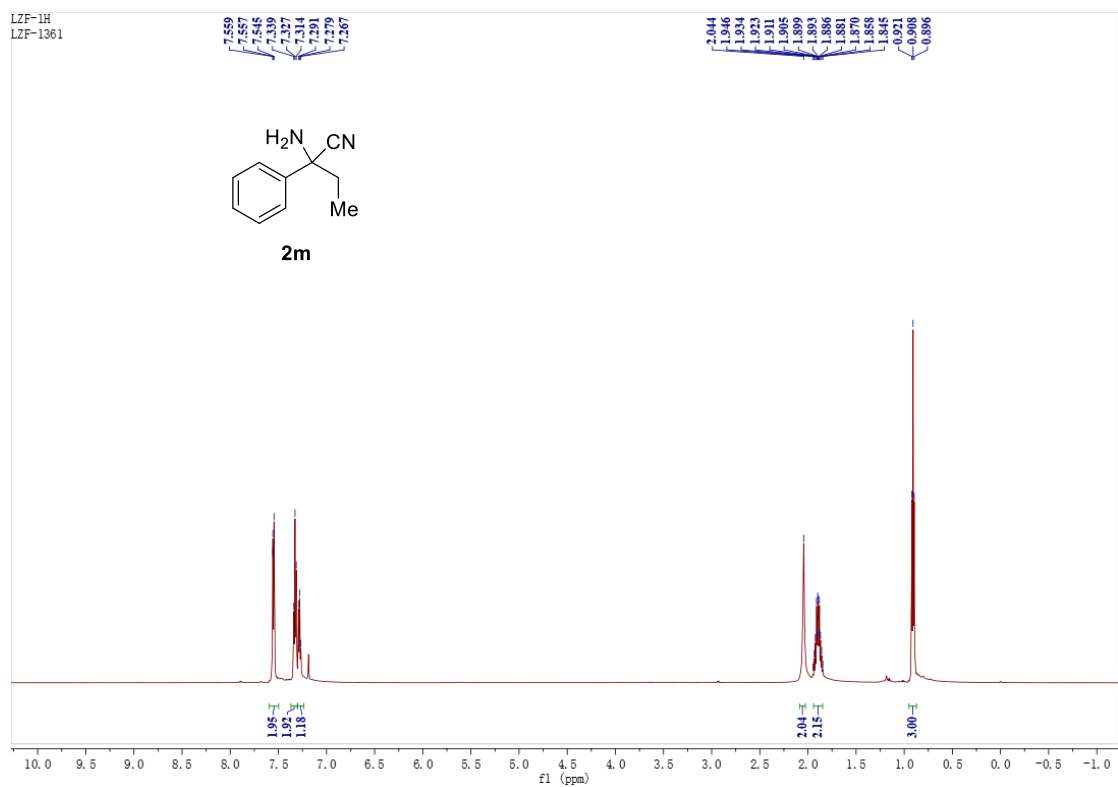




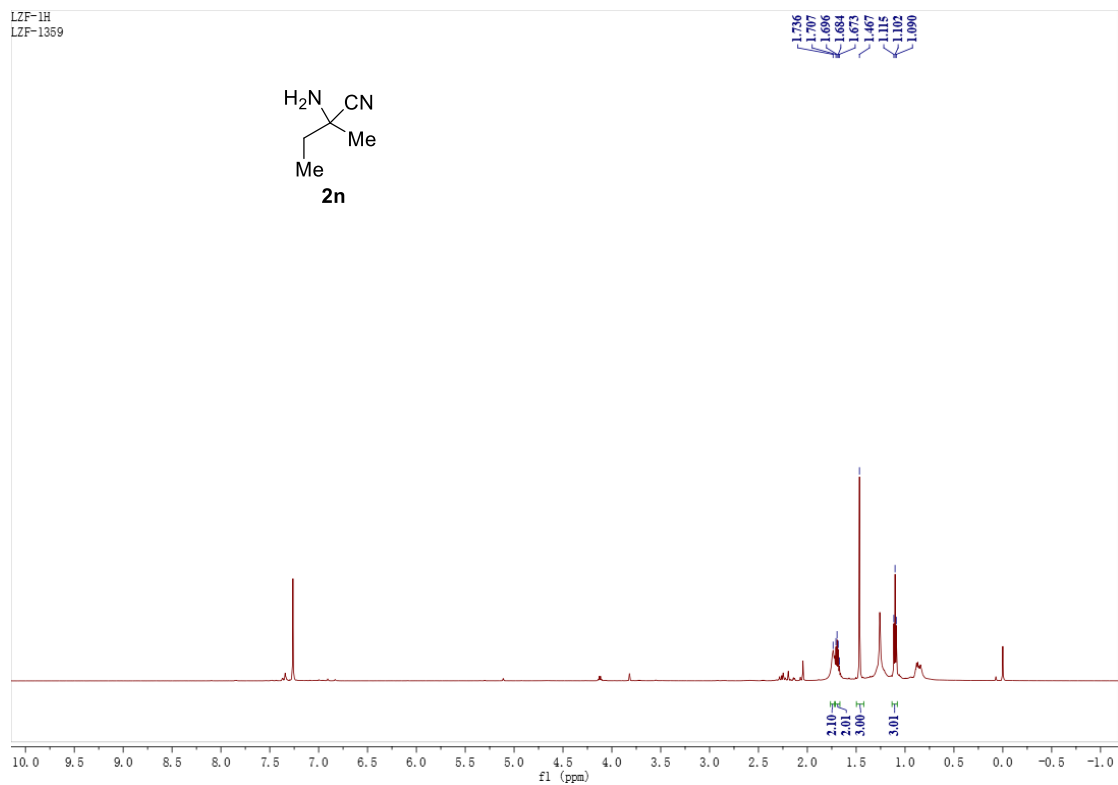
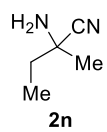




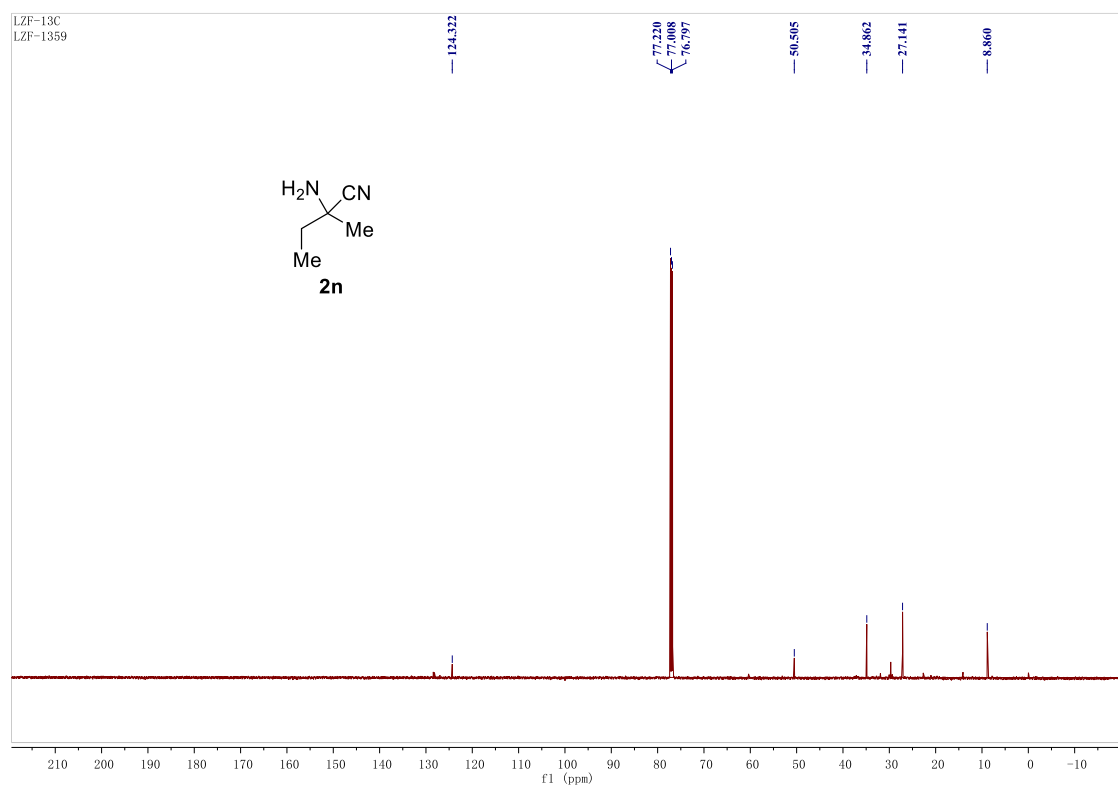
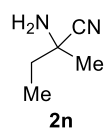


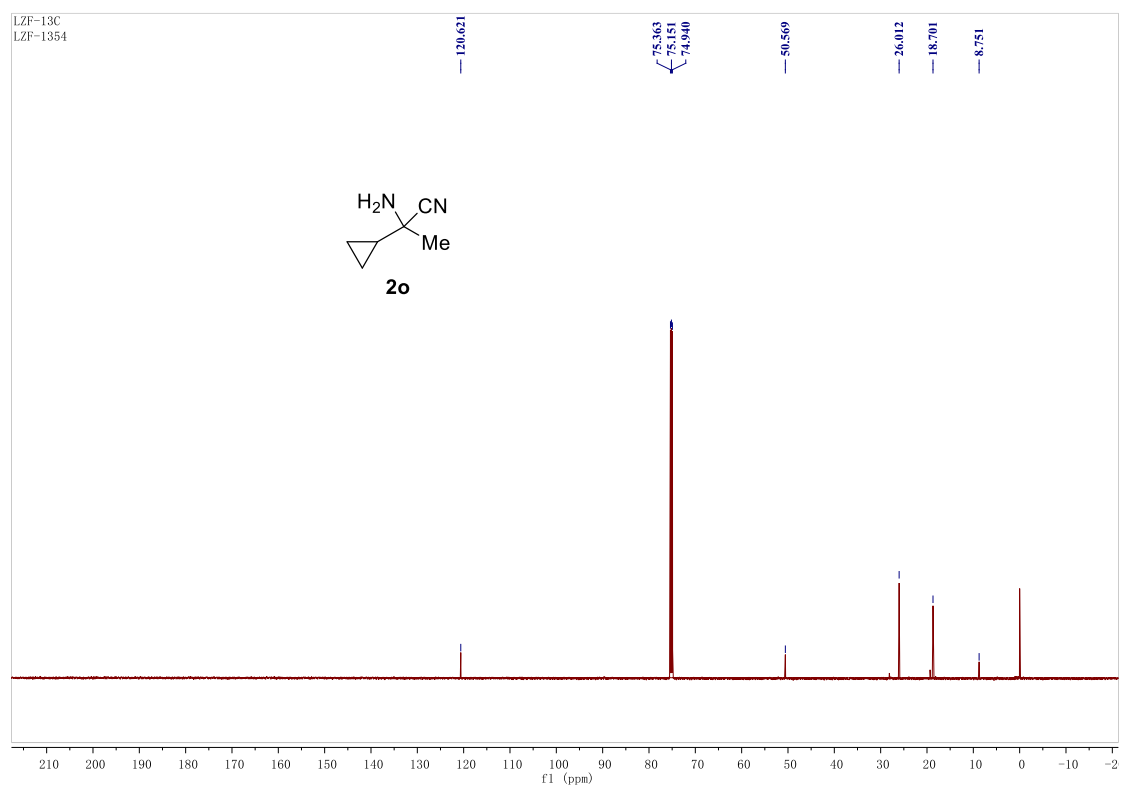
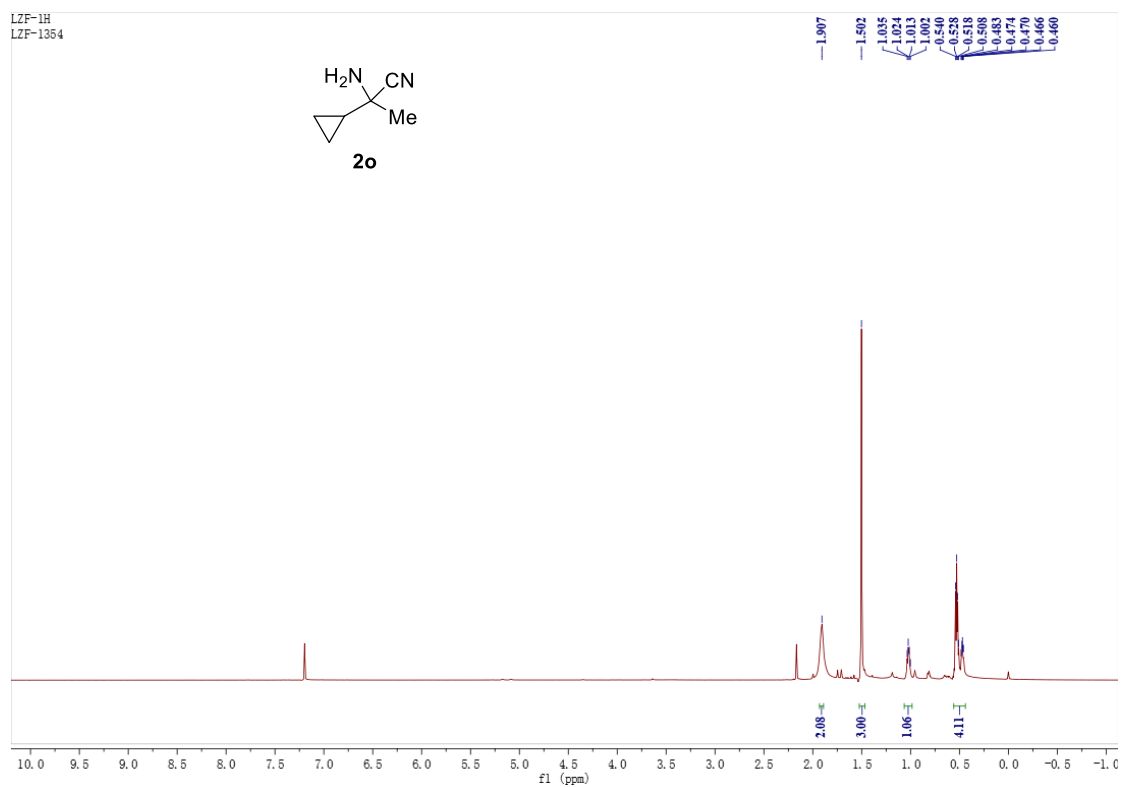


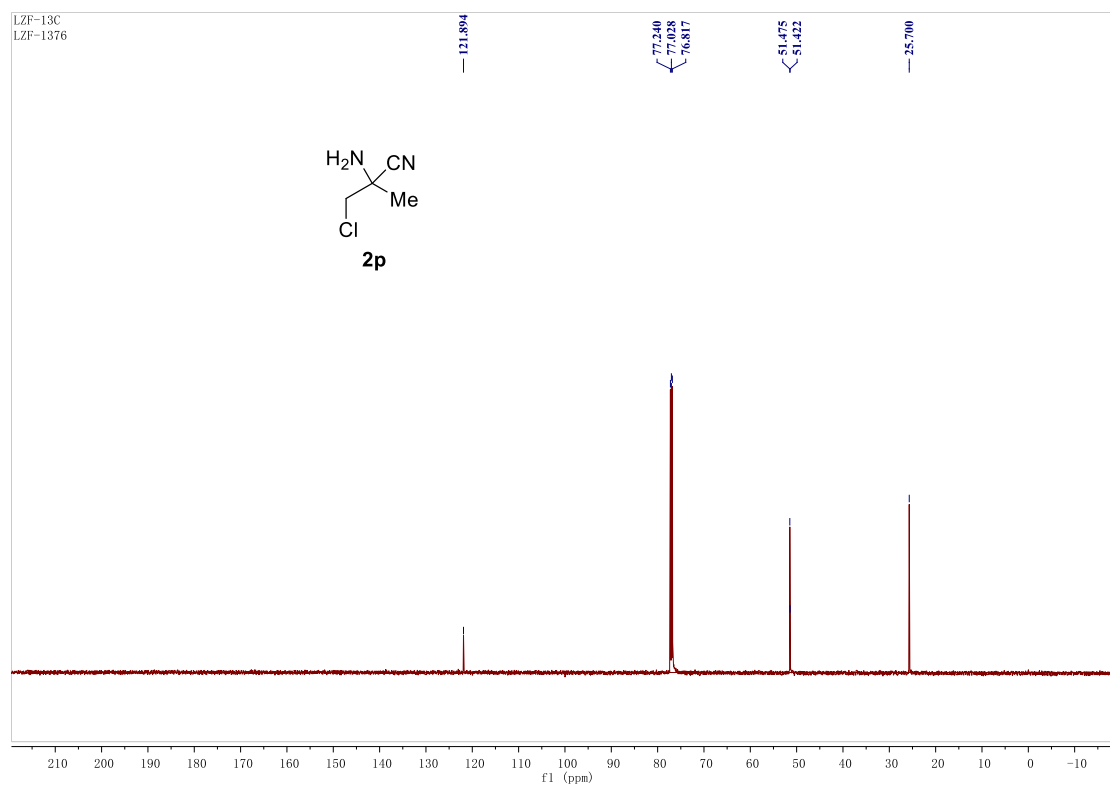
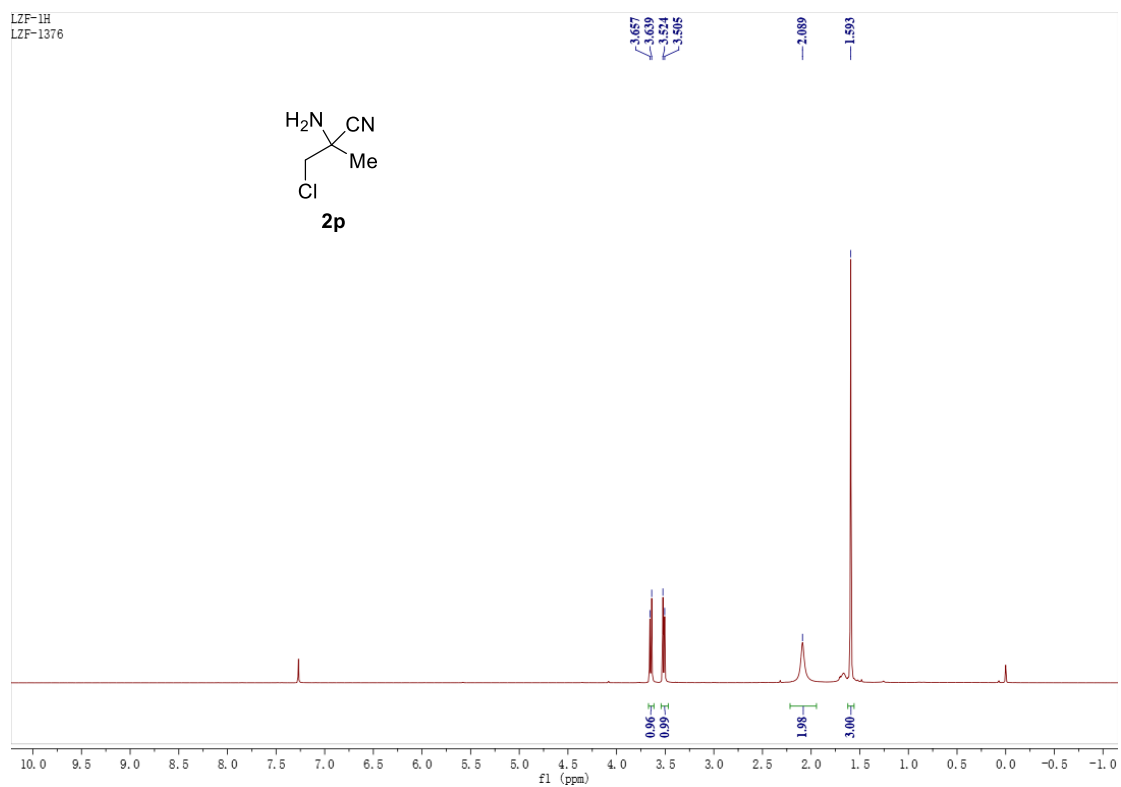
LZF-1H
LZF-1359

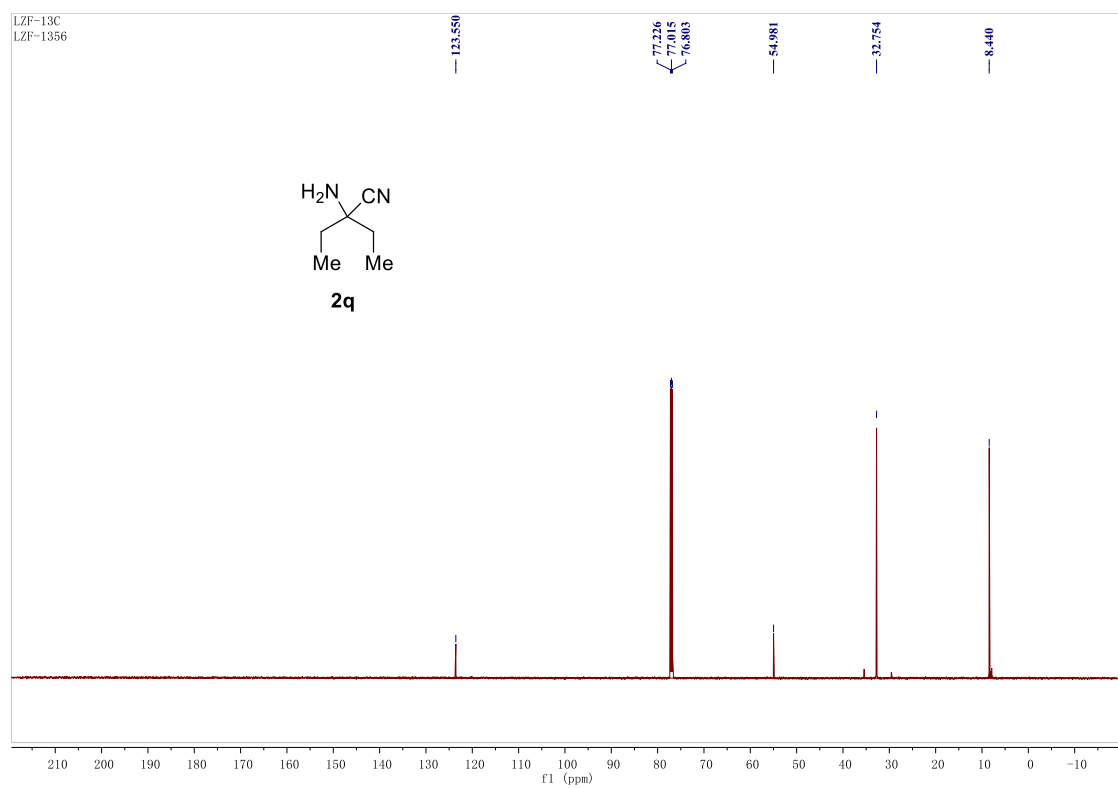
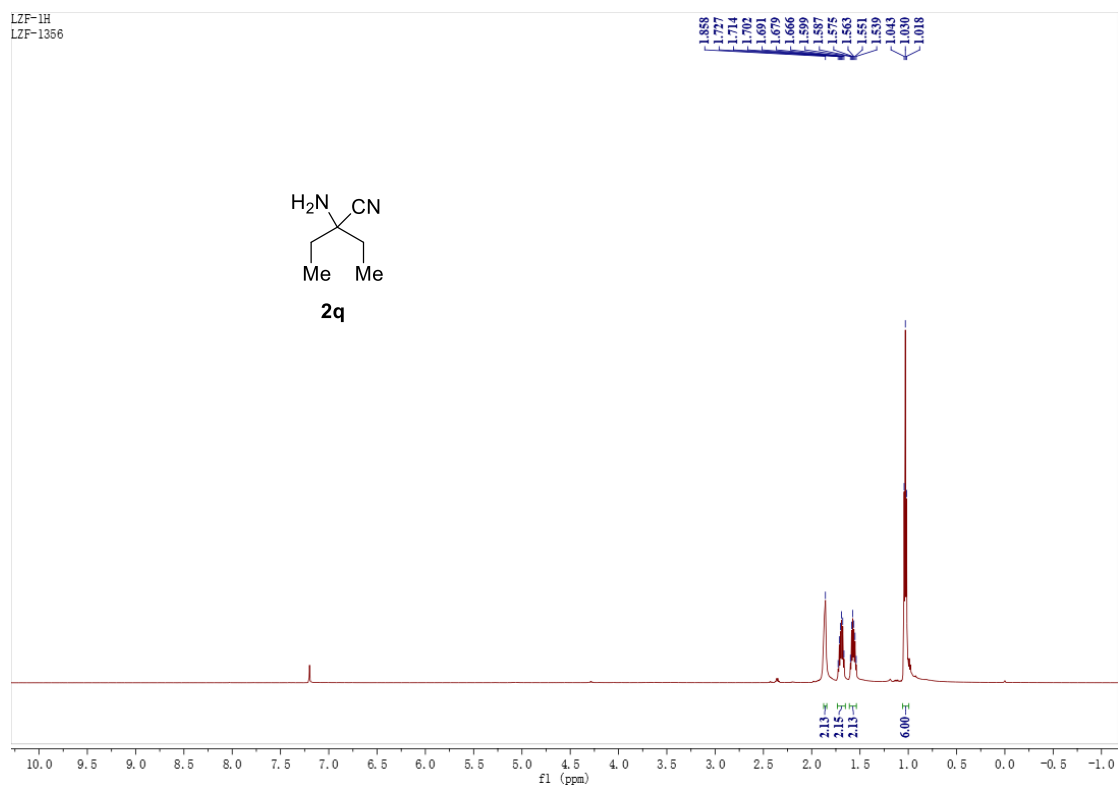


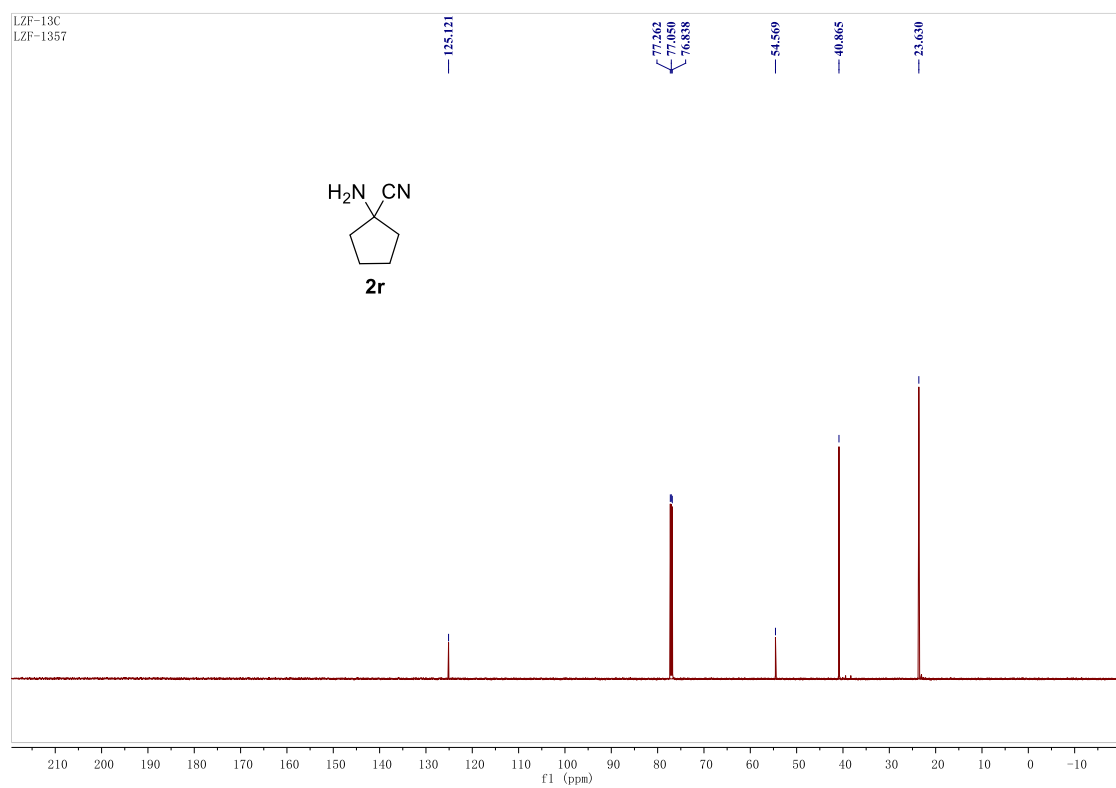
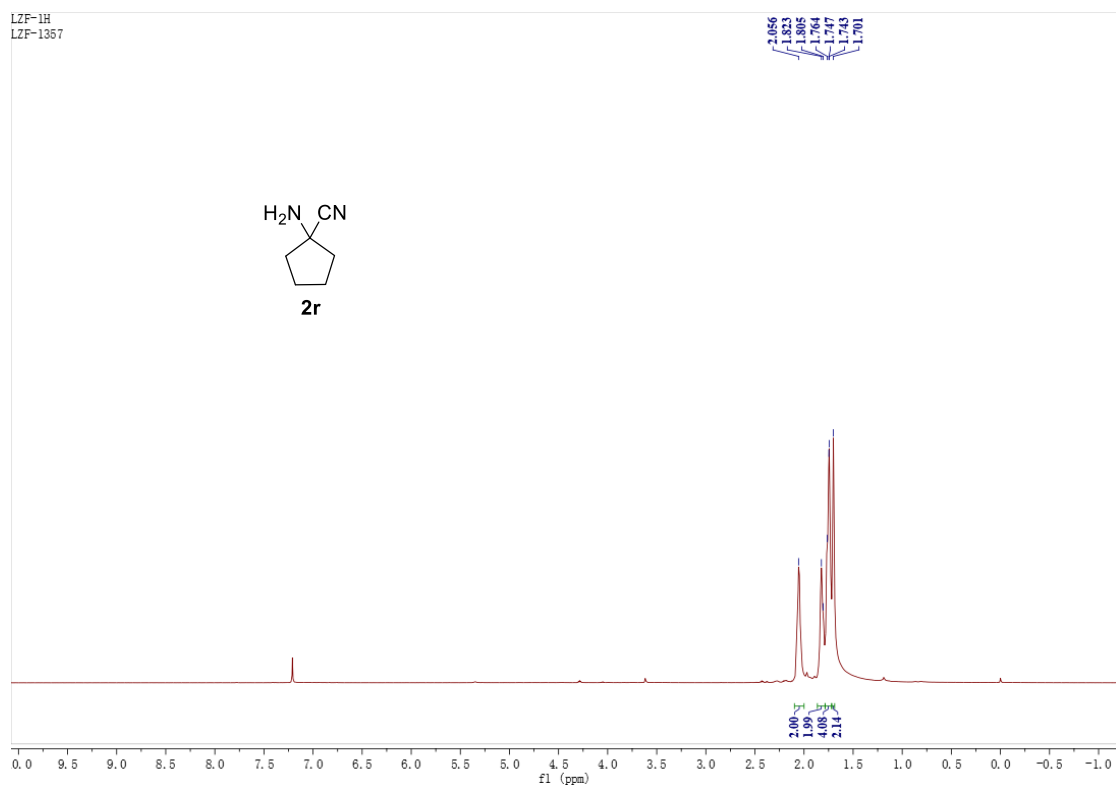
LZF-13C
LZF-1359

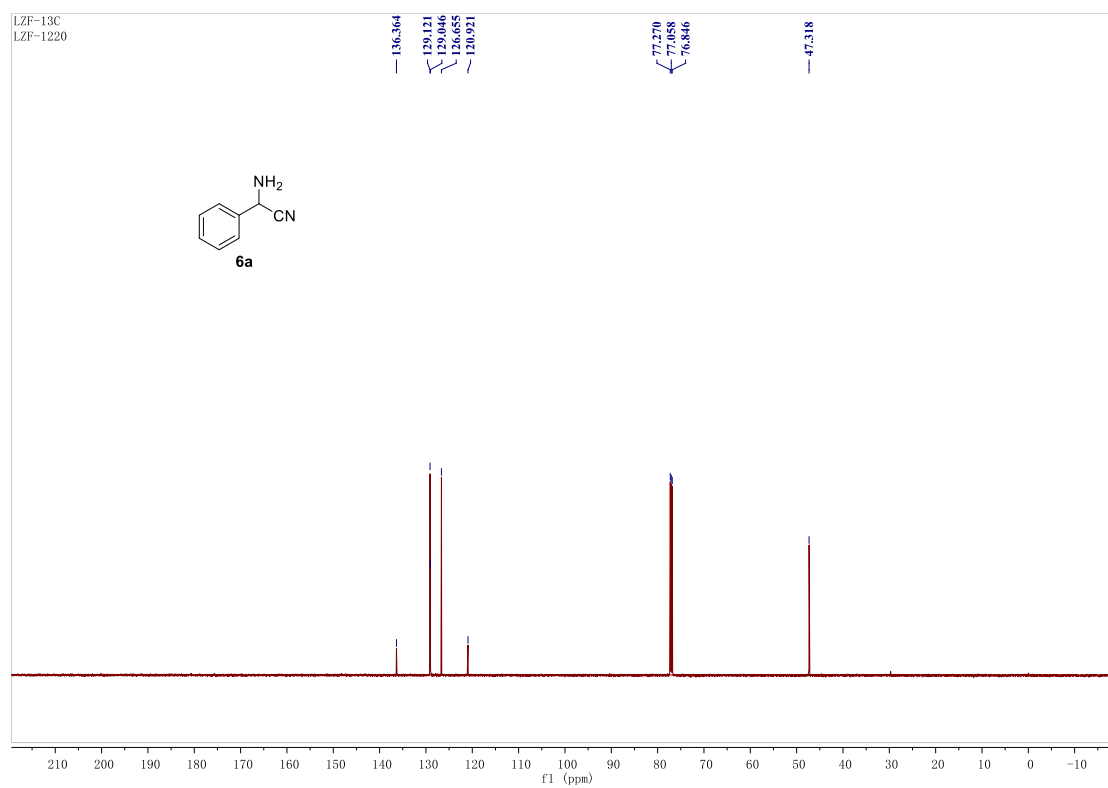
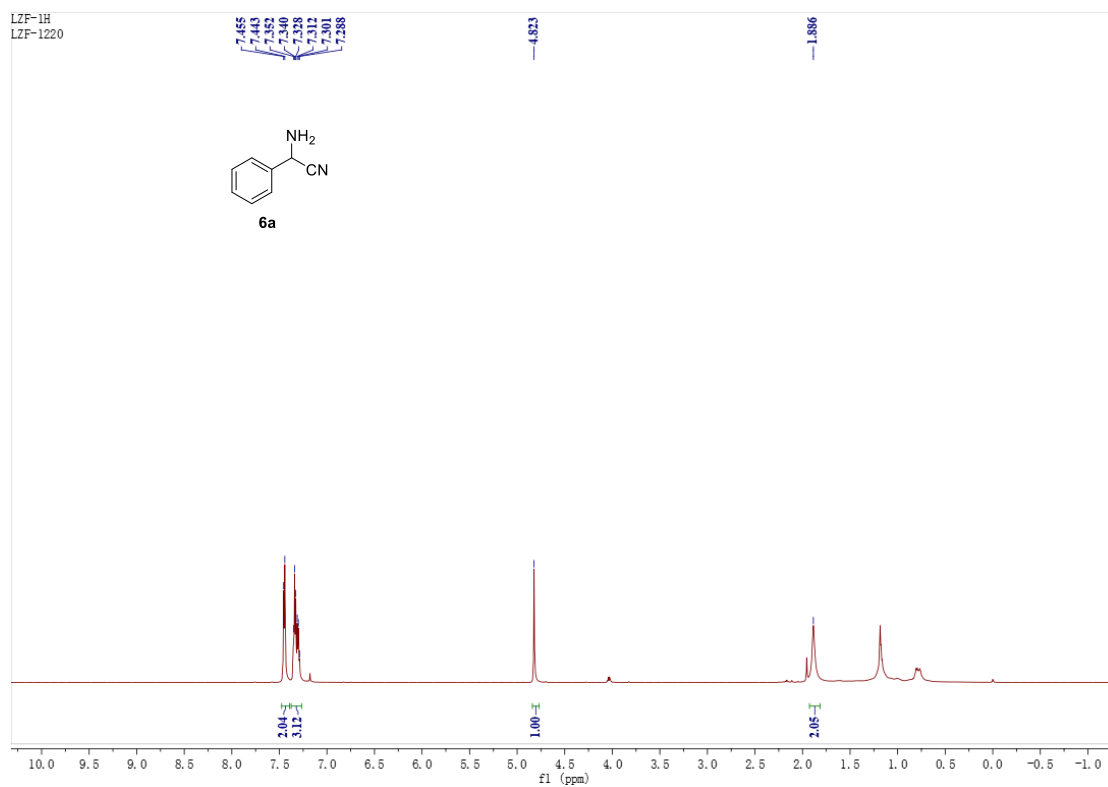


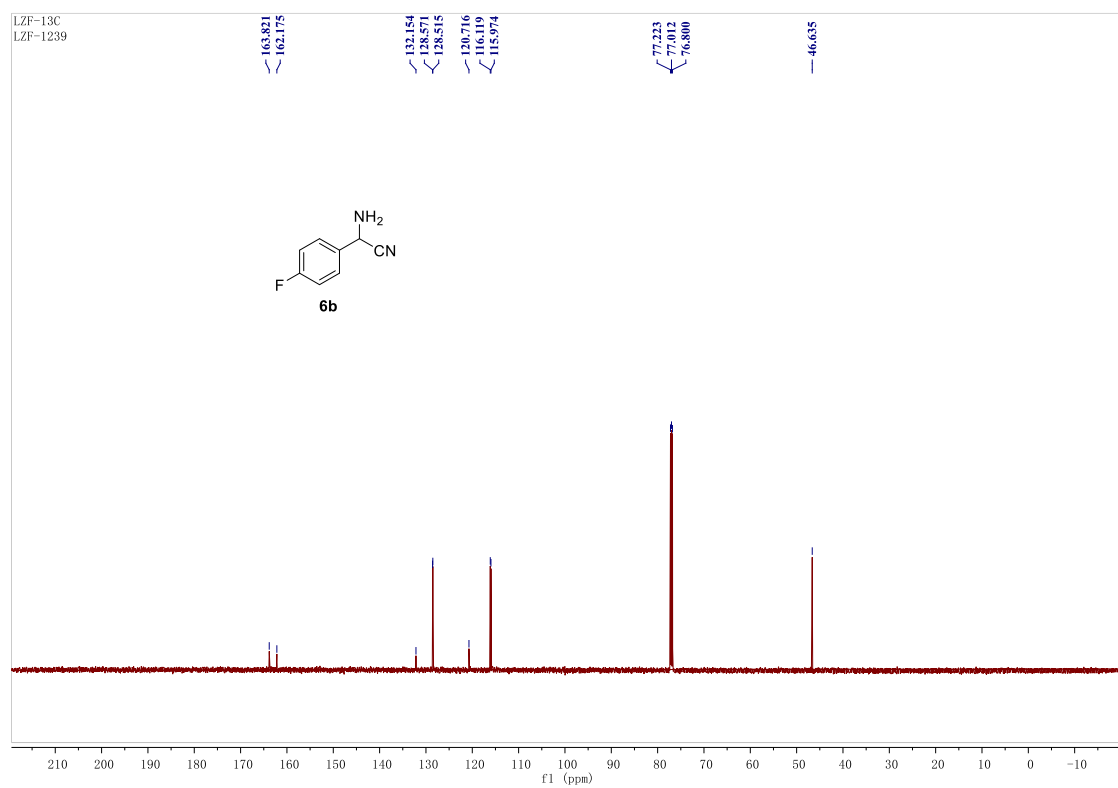
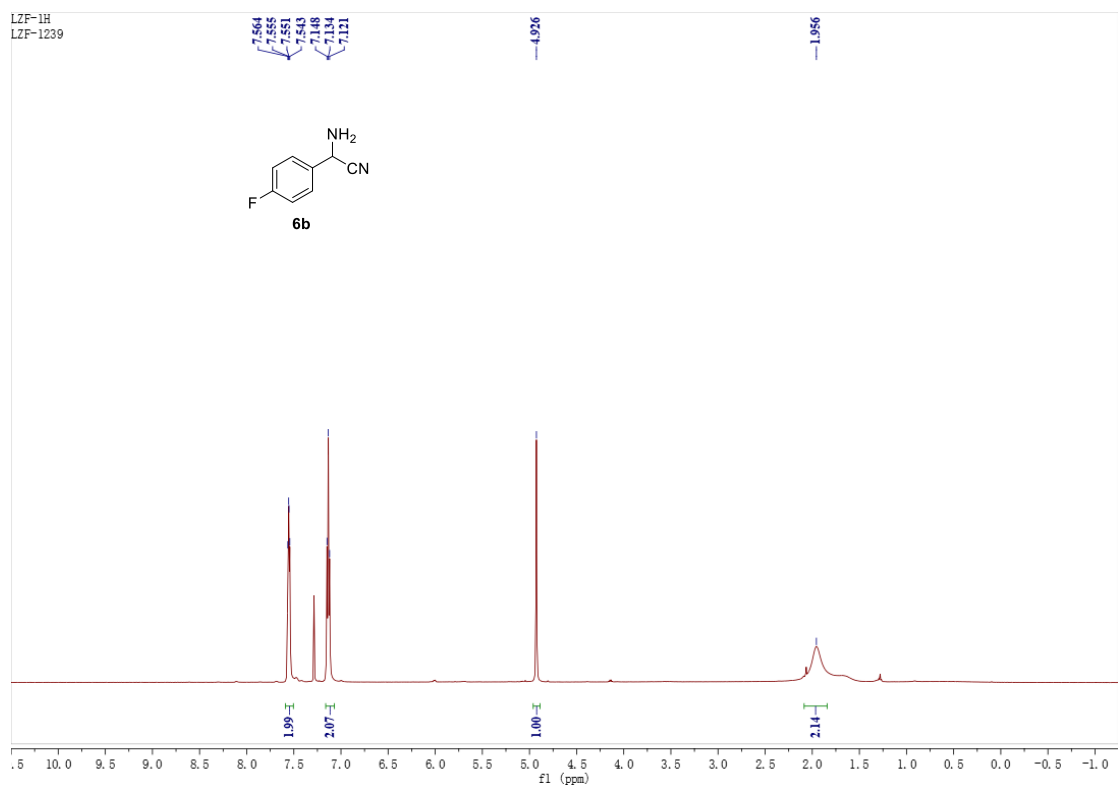


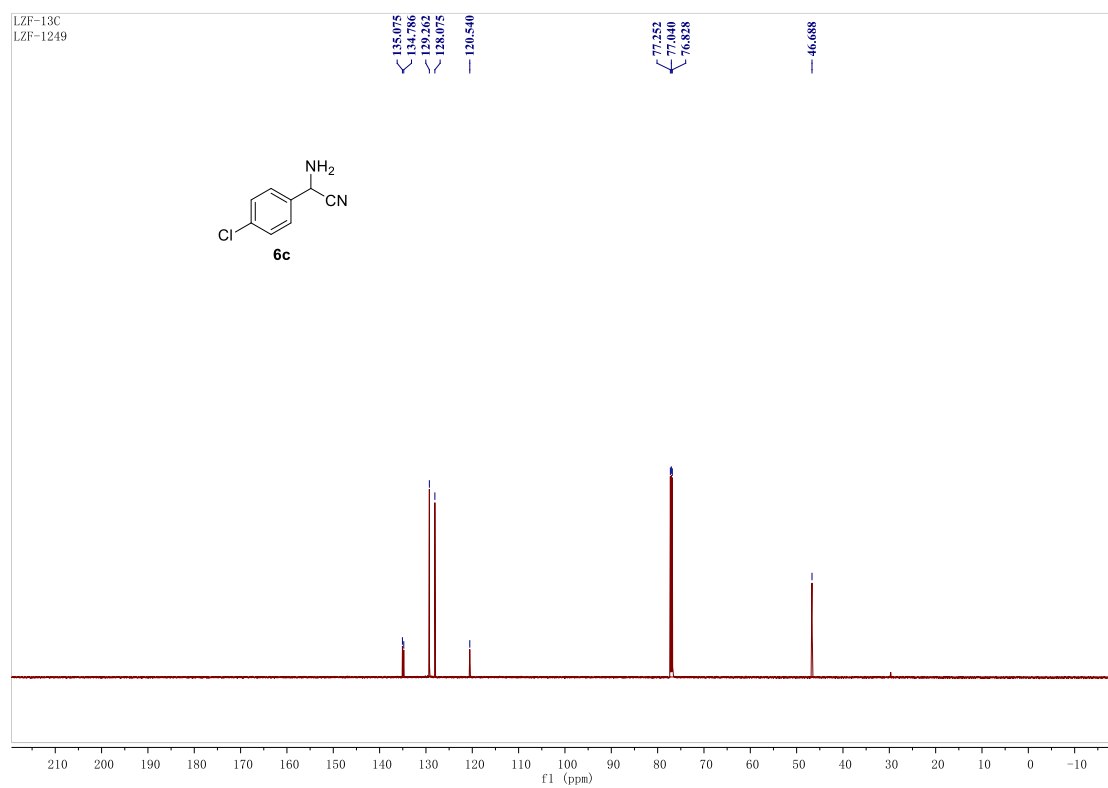
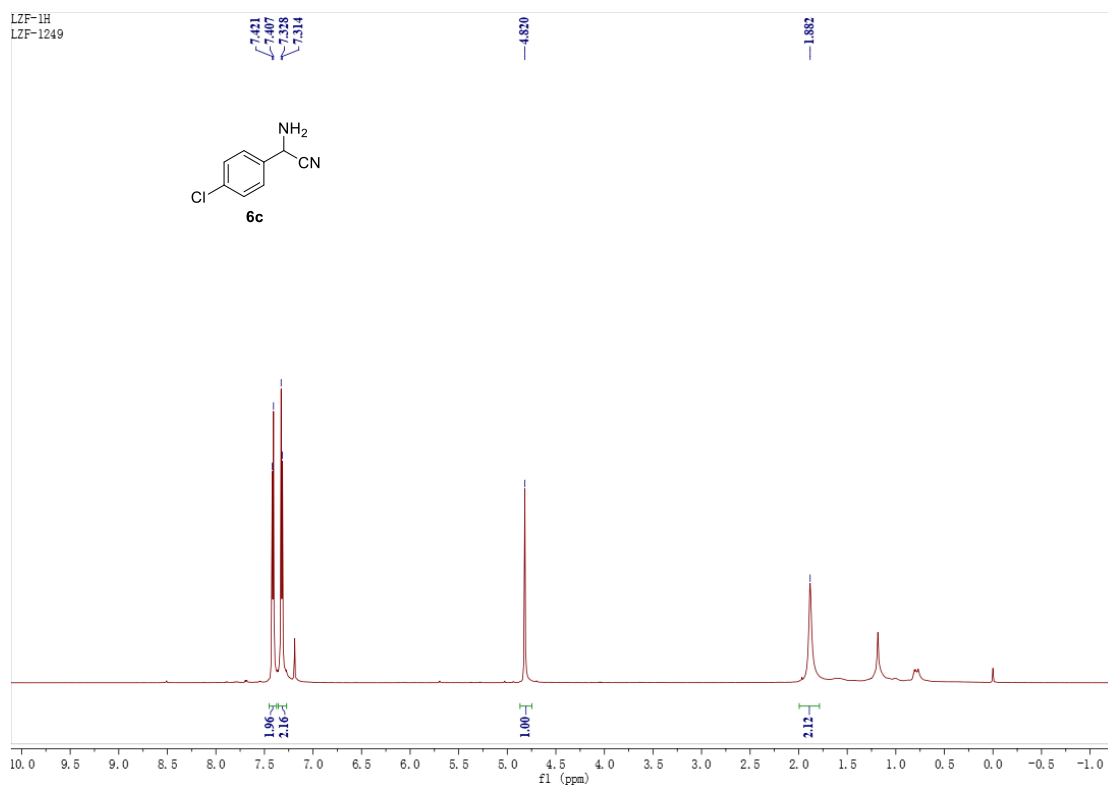


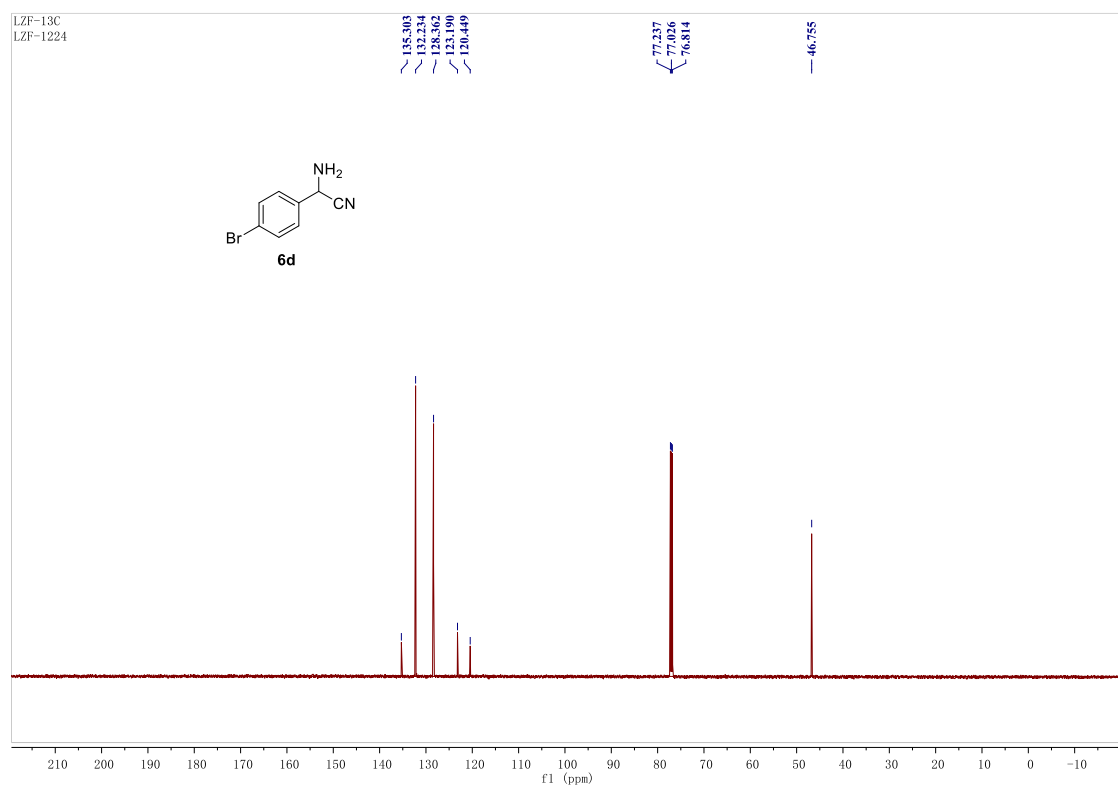
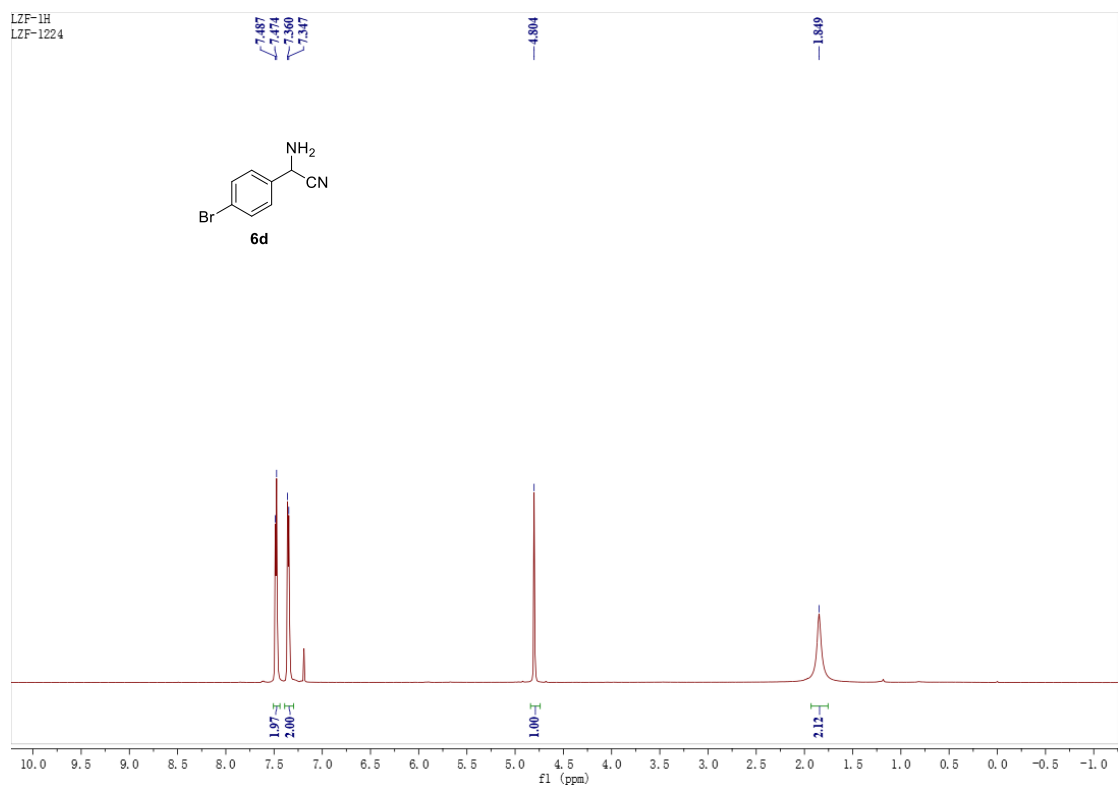


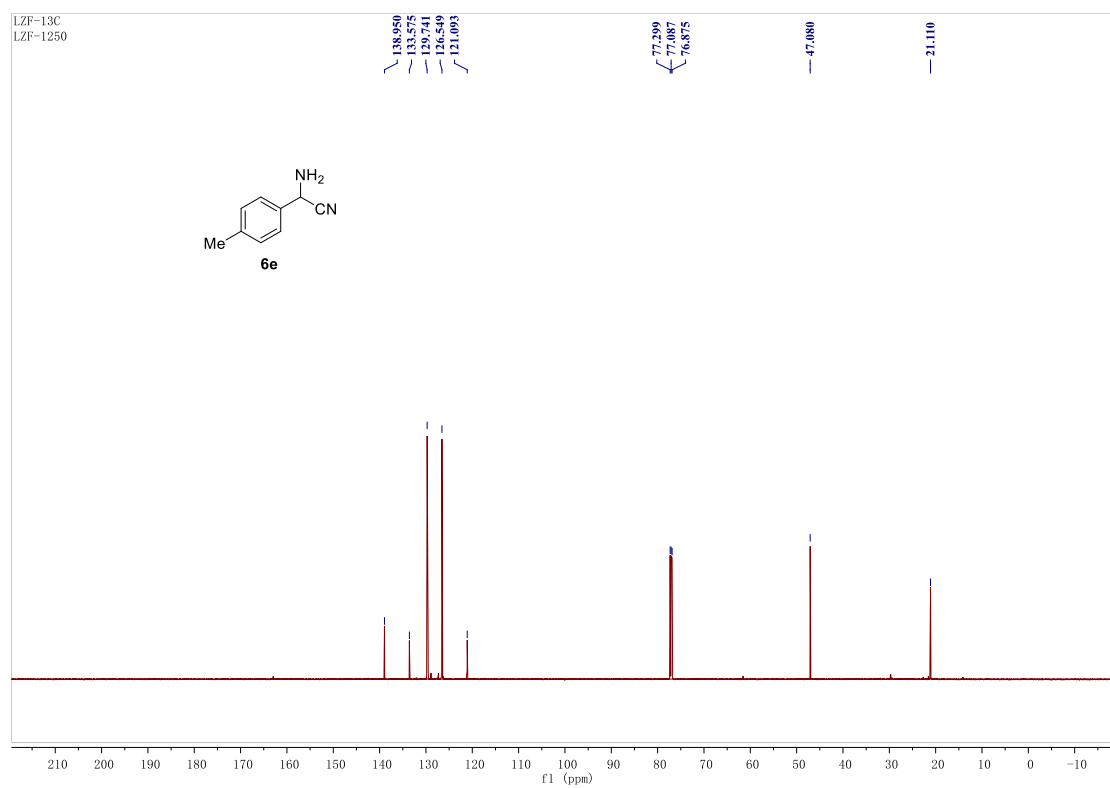
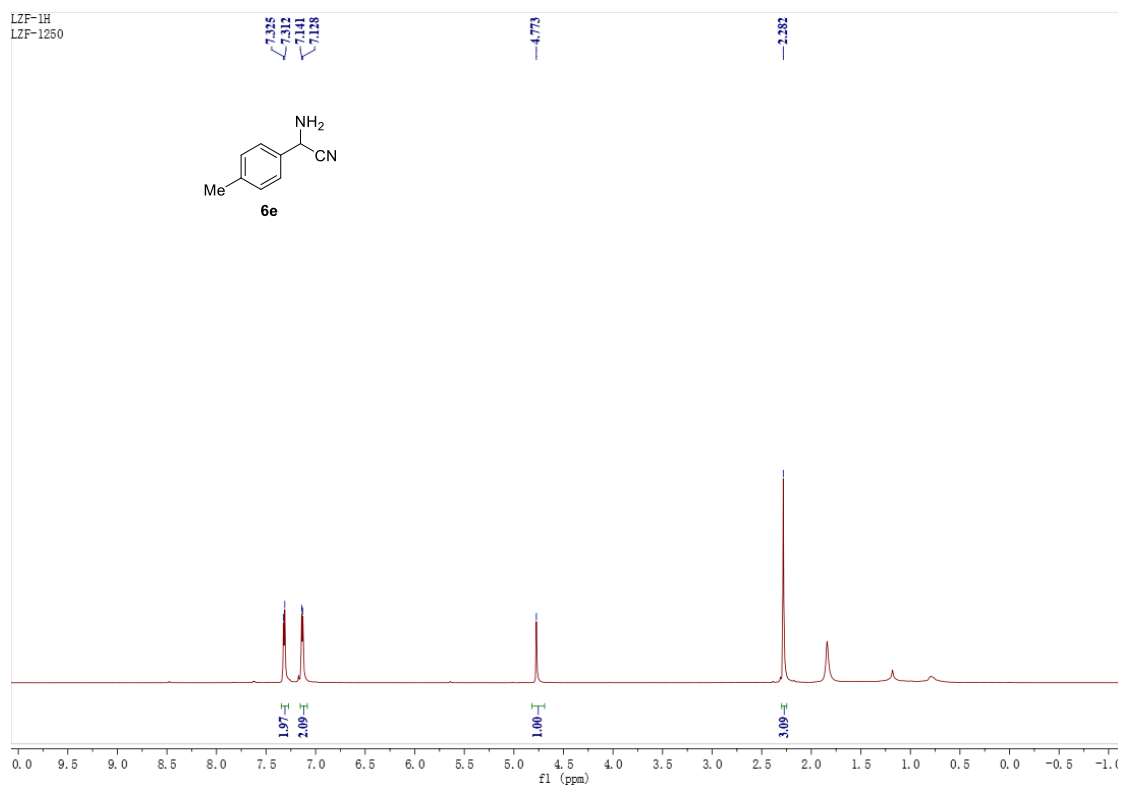


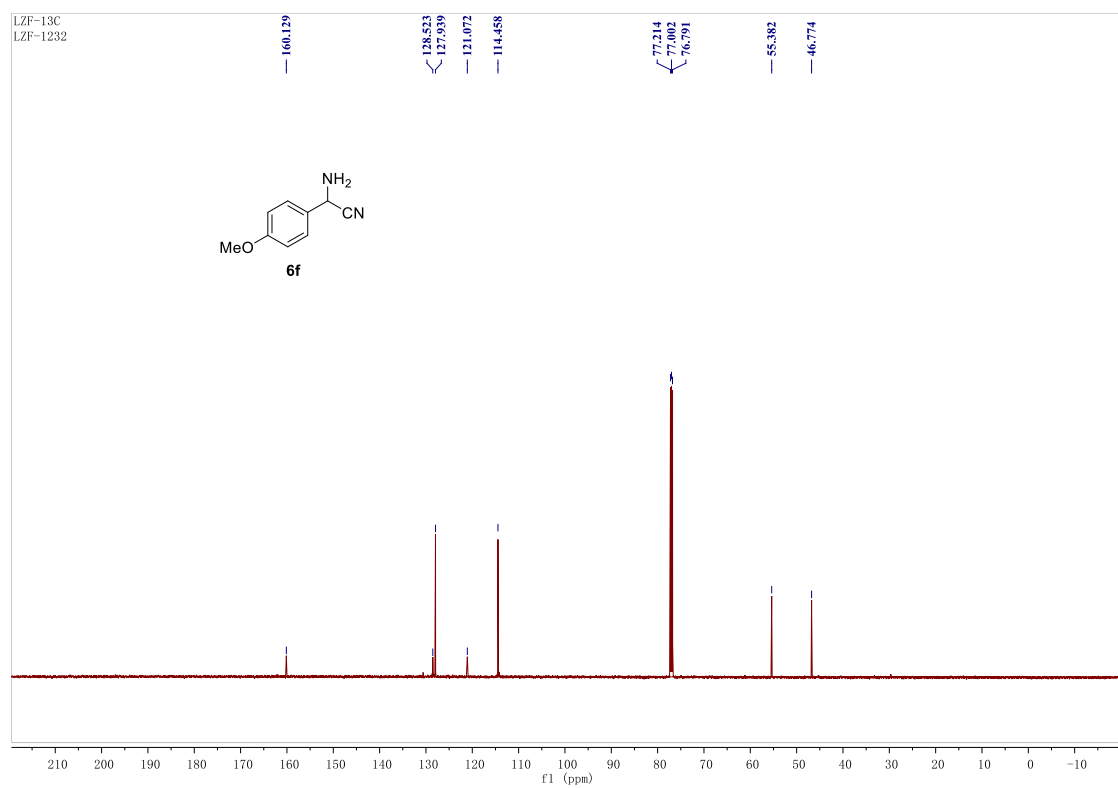
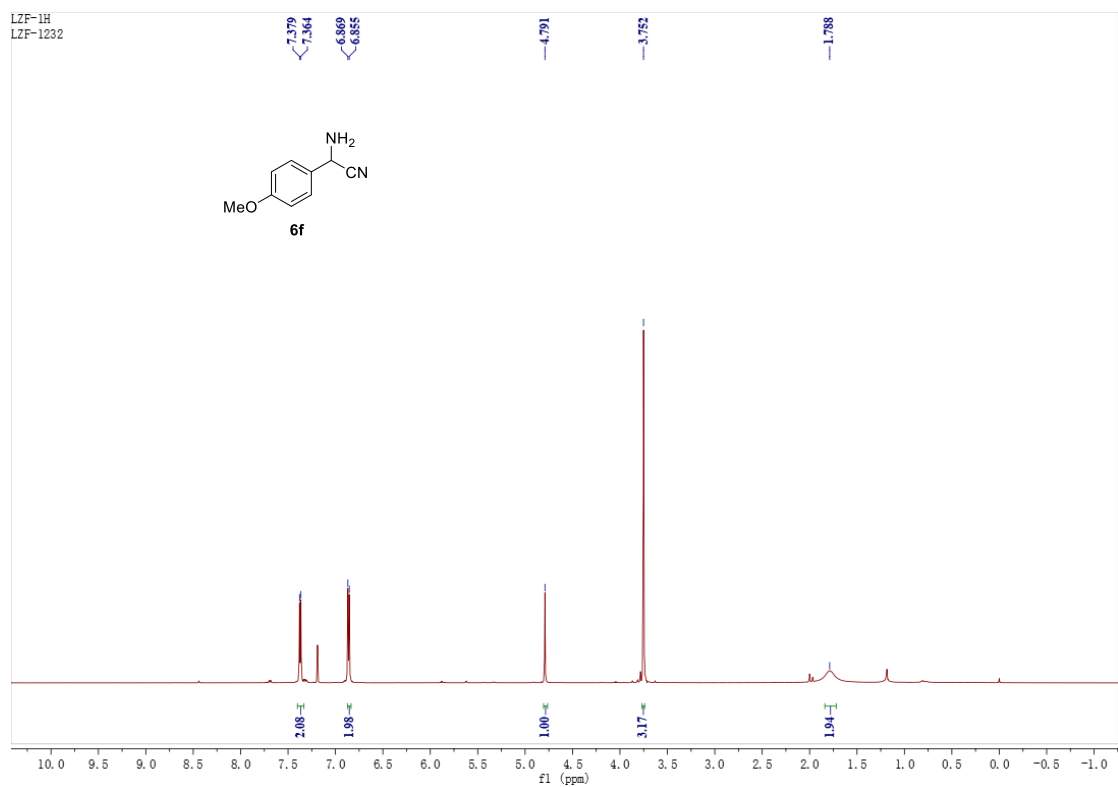


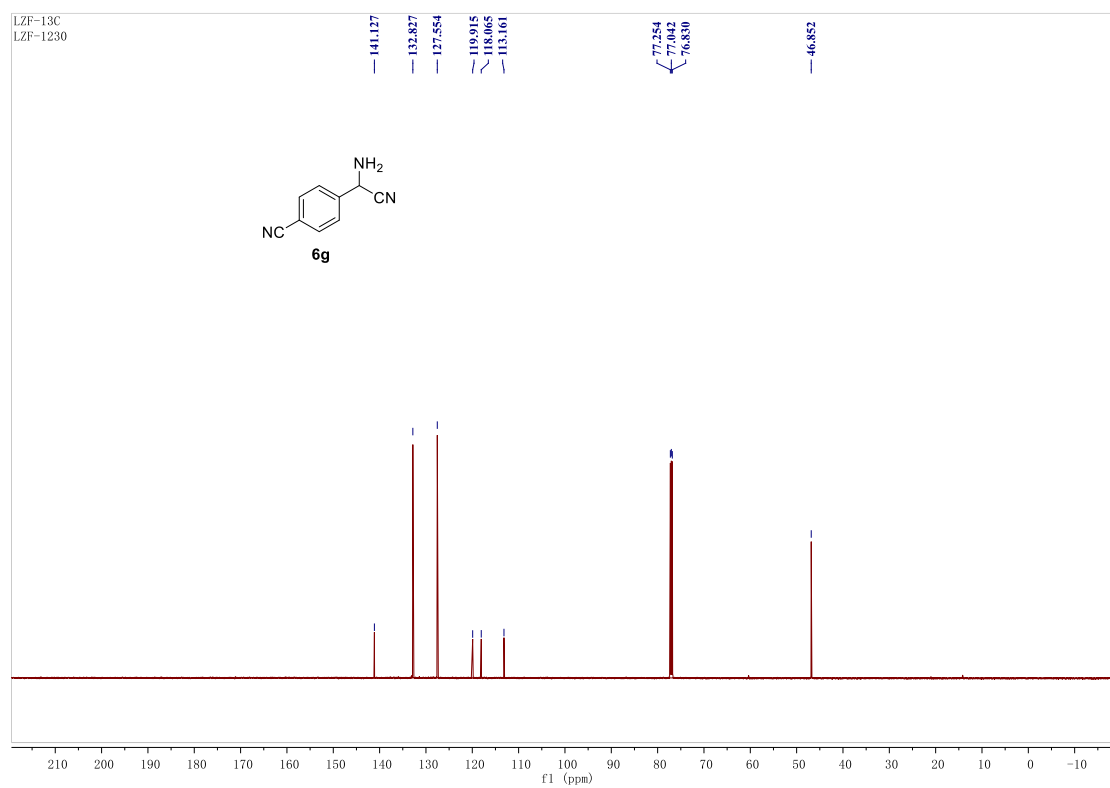
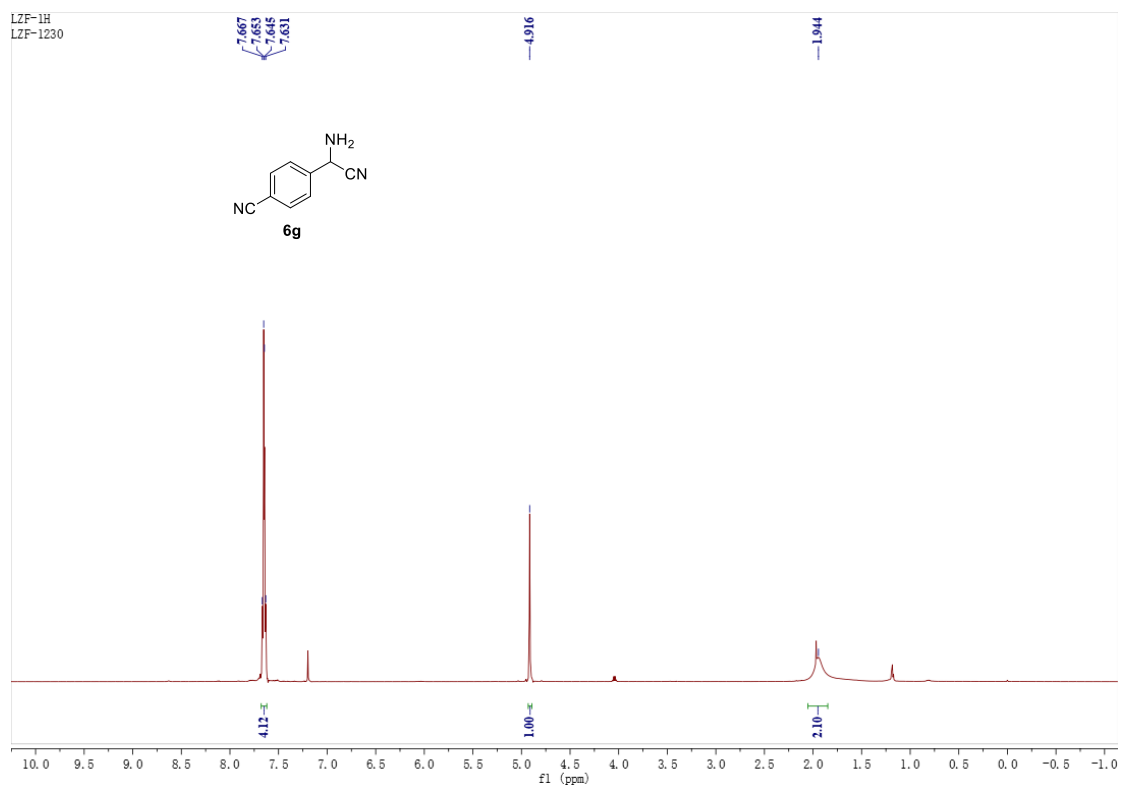


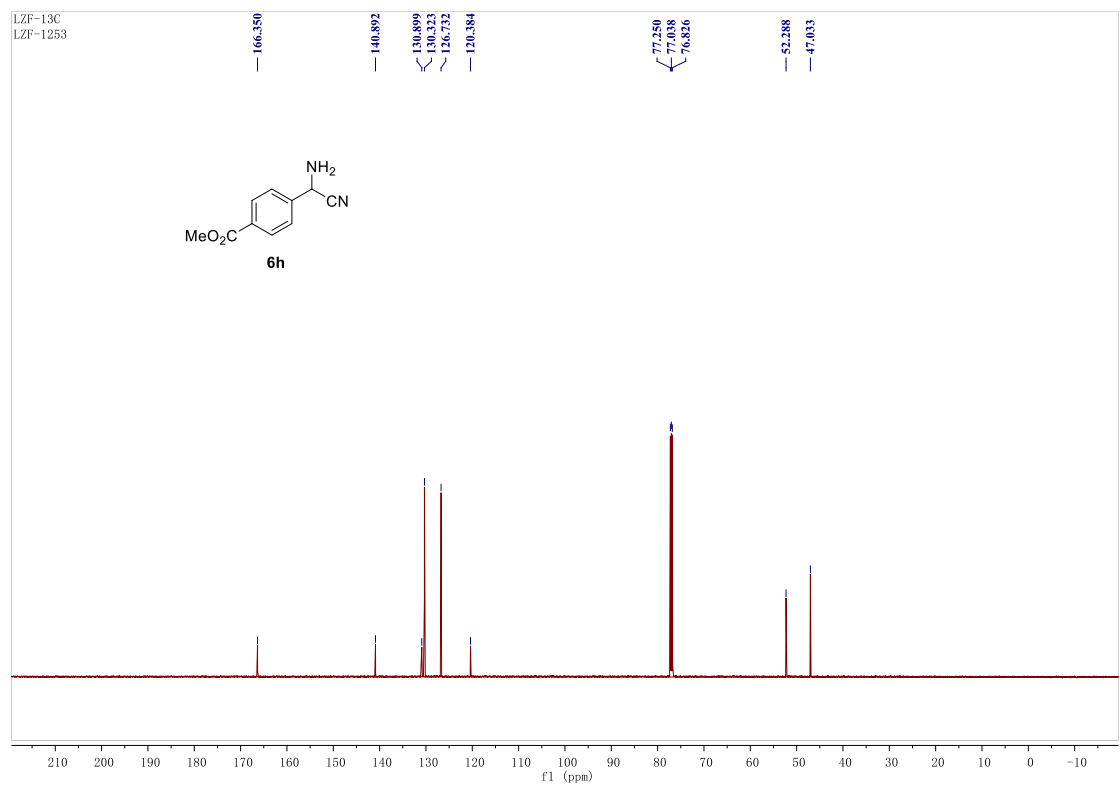
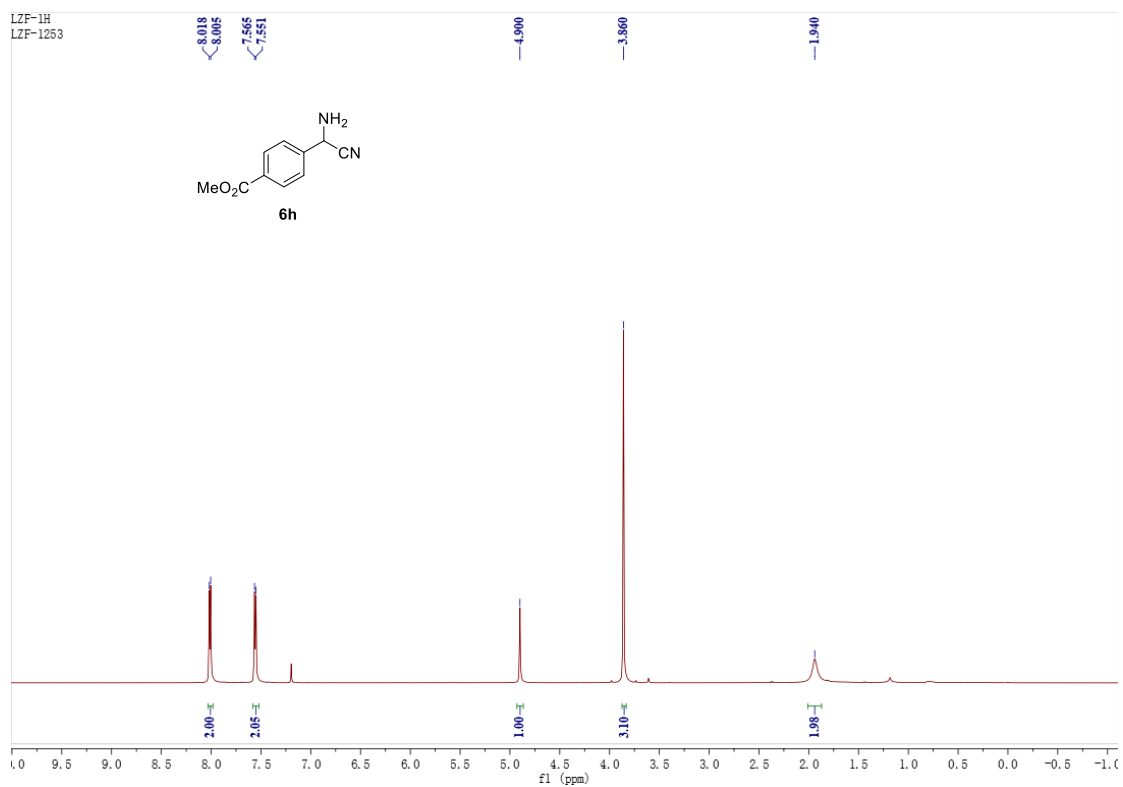


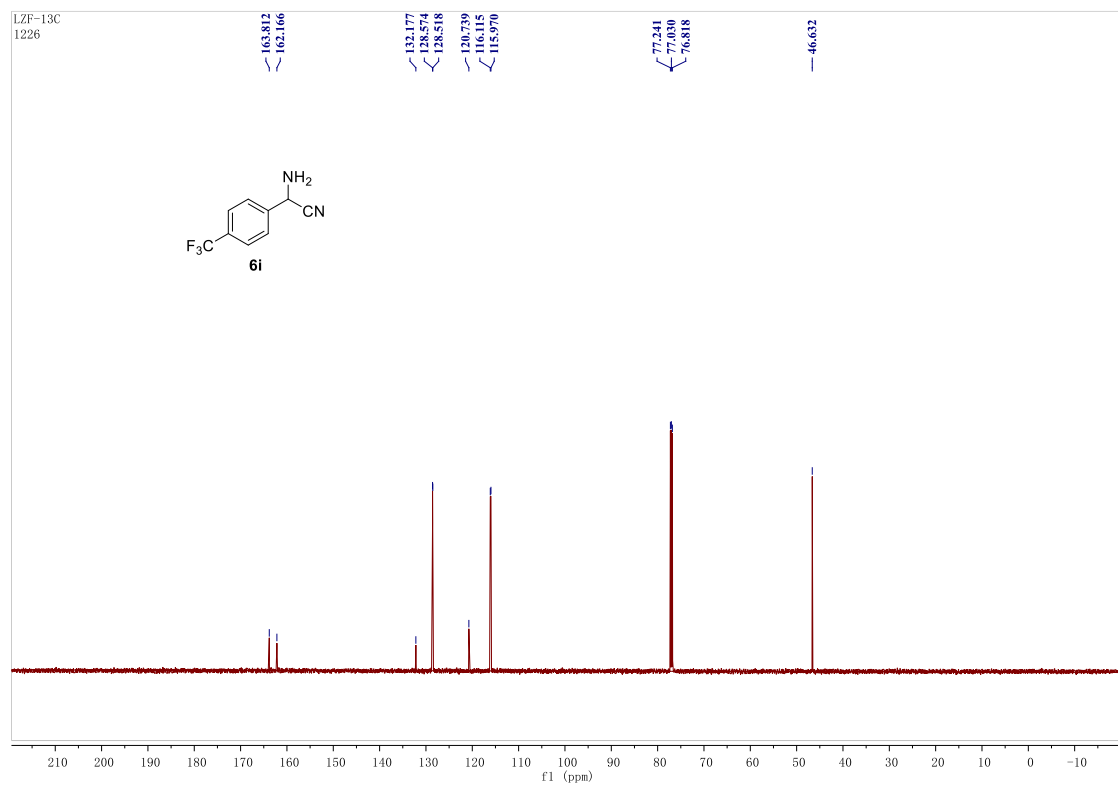
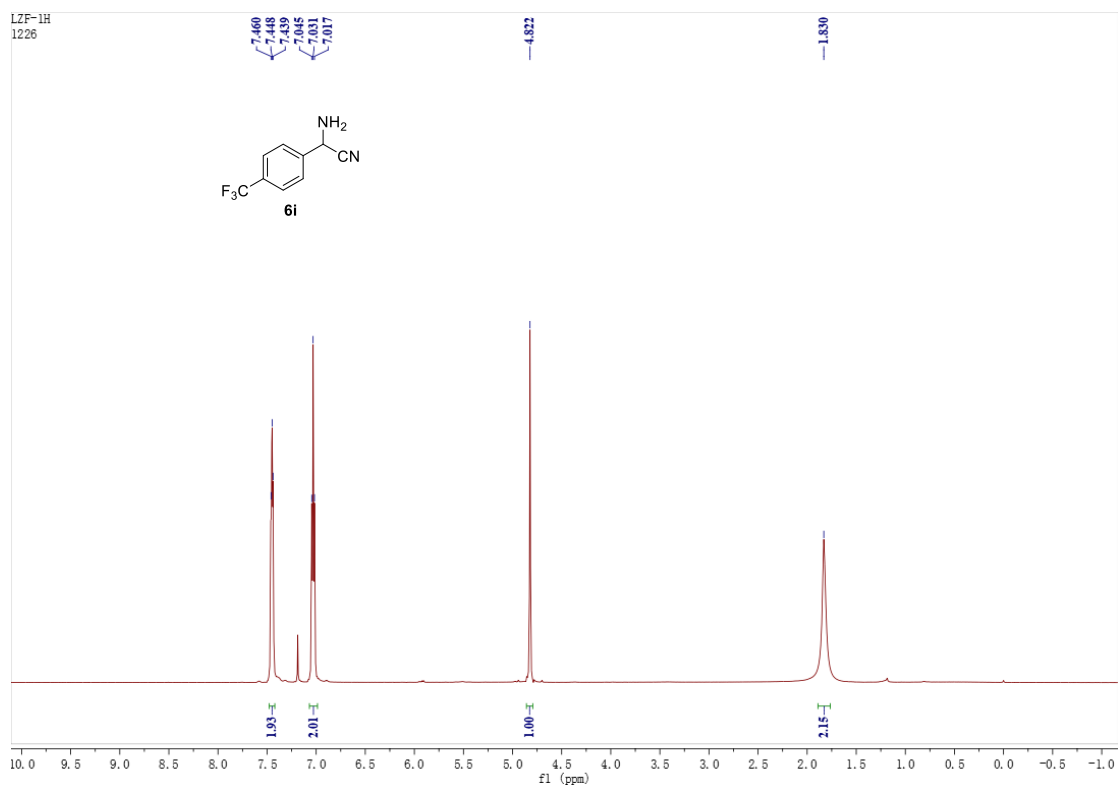


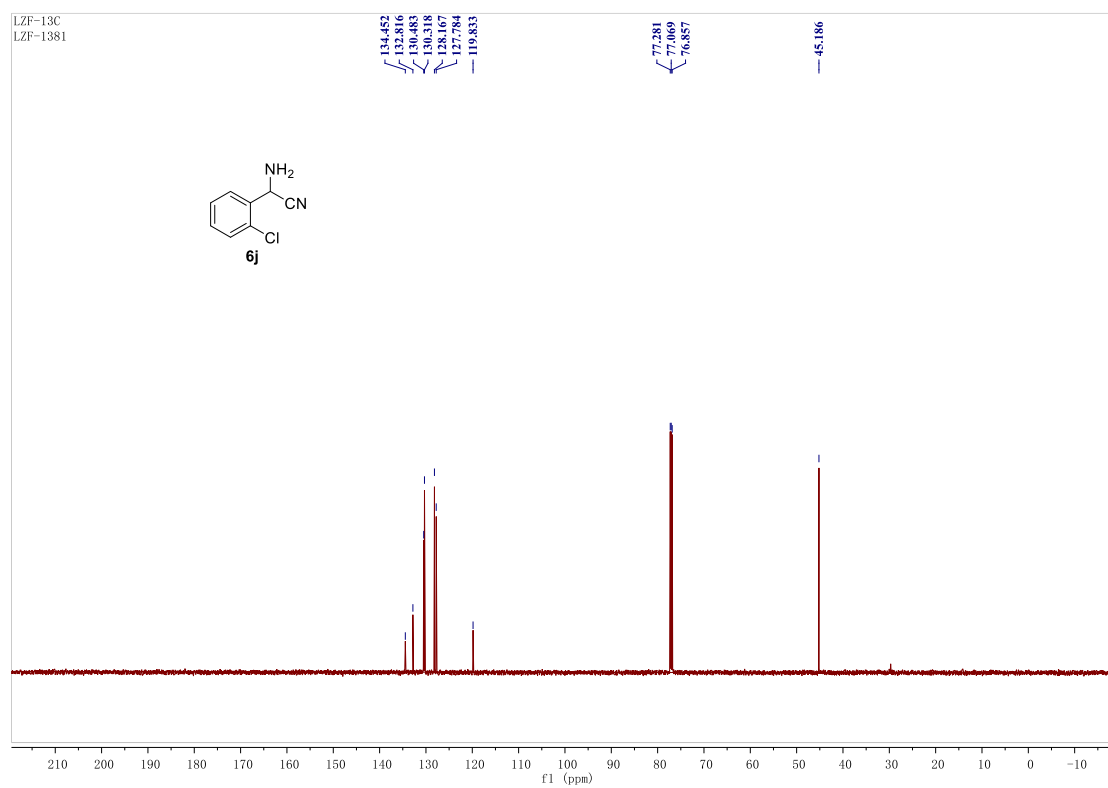
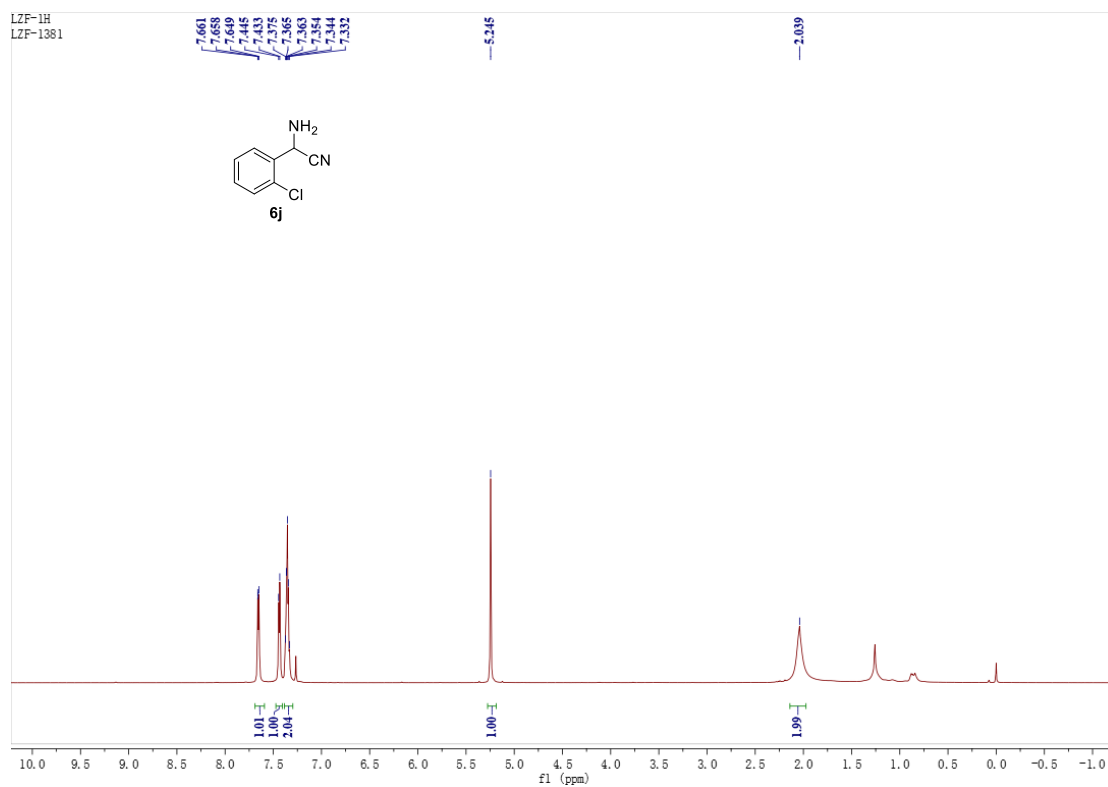


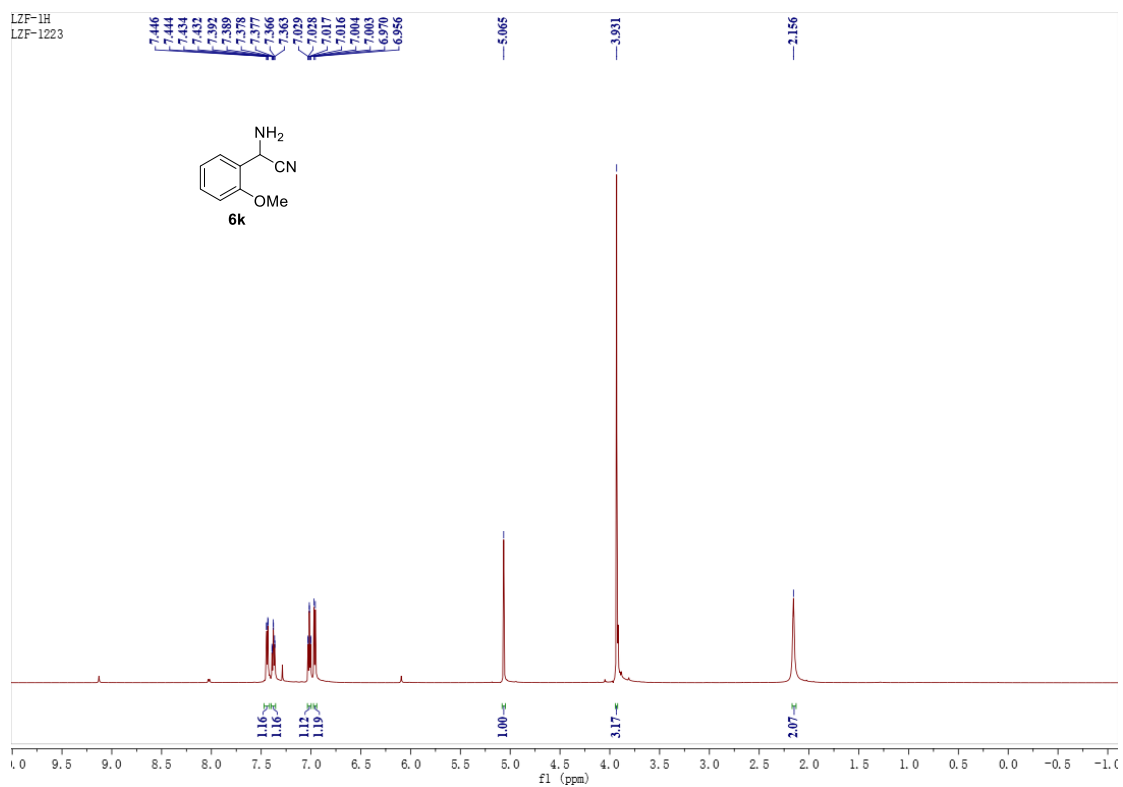












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