

Supporting information

Solvent and catalyst-free tandem reaction: Synthesis, photophysical and biological application of isoindoloquinazolinones

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General information:

^1H NMR (400 MHz) and ^1H NMR (500 MHz) spectra were recorded on a 400 MHz and 500 MHz spectrometer in CDCl_3 / $\text{DMSO}-d^6$ solvent using TMS as the internal standard. ^{13}C NMR was recorded in the same MHz instruments. HRMS was measured using a TOF analyzer. 60-120 or 100-200 mesh silica gels (SRL) were used for column chromatographic purification. Progress of the reaction was monitored by using precoated silica gel 60 F254 TLC sheets (Merck). Petroleum ether (boiling range 60-80 °C) or n-hexane was used as the eluent for column chromatographic separation. Solvents were distilled, dried and stored over molecular sieves (4 Å). The UV-Vis absorption spectra were recorded on a Shimadzu UV-2450 UV-Vis spectrophotometer and the fluorescence emission spectra were recorded on Photon Technology International S/N-3201.

General Procedure for the preparation of 1a – 1j:¹

β -Bromovinyl aldehyde or 2-Bromo Benzaldehyde **1** (1 mmol), Na_2CO_3 (4 mmol), Bu_4NBr (1 mmol), PdCl_2 (10 mol%), and H_2O (5 mL) were taken in a two-neck, round-bottom flask. Acrylic ester (4 mmol) was added, and the mixture stirred for 2 h in 50 °C, diluted with brine solution, and extracted with EtOAc ($3 \times 25\text{mL}$). The extracted solution was dried over Na_2SO_4 , and the solvent was evaporated. The product was purified by column chromatography using EtOAc–PE as eluent.

Preparation of (E)-3-(2-formylcyclohept-1-en-1-yl)acrylonitrile (1k):

2-bromocyclohept-1-ene-1-carbaldehyde (1 mmol), $\text{P}(\text{o-tolyl})_3$ (0.25 mmol), $\text{Pd}(\text{OAc})_2$ (10 mol%), triethyl amine (1mL), methanol (1mL), acetonitrile (1mL) were placed in a two-neck, round-bottom flask. Acrylonitrile (2 mmol) was added and the mixture stirred in inert atmosphere for 2 hr in 80 °C and extracted with EtOAc ($3 \times 25\text{ mL}$). The extracted solution was dried over Na_2SO_4 , and the solvent was evaporated. The product was purified by column chromatography using EtOAc–PE as eluent.

Preparation of 2-amino-5-nitrobenzamide (2e):

Isatoic anhydride (1 mmol) was dissolved in 5mL conc. H_2SO_4 and it was placed in ice bath. NaNO_3 (1 mmol) was taken in another round-bottom flask and it also kept in ice bath. The acidic solution of Isatoic anhydride was added drop wise to the ice cooled NaNO_3 and the resulting mixture stirred continuously for 1 hr. Finally the reaction mixture was poured in ice cube , a yellow ppt. was obtained. This ppt. was collected by filtration and dried.

This yellow ppt. was dissolved in 15 mL liq. NH_3 . This mixture was allowed to stir in reflux condition at $\sim 80^\circ\text{C}$ for 8 hr. Finally this reaction mixture was recovered by filtration. Yellow ppt. was collected and dried.

General Procedure for preparation of (3a – 3t):

Compound **1** (1.0 equiv.) and **2** (1.0 equiv.) were taken in a round-bottom flask and it was placed in a heating oil bath. The heating of the resulting mixture was carried out up to 2 h. in open flask at 120°C under the neat condition. The progress of the reaction was monitored by TLC. The crude residue was purified by silica-gel (100-200 mesh) column chromatography using ethyl acetate and petroleum ether (1:2) as an eluent to get the desired product **3**.

UV/Vis absorption spectra of **3j**:

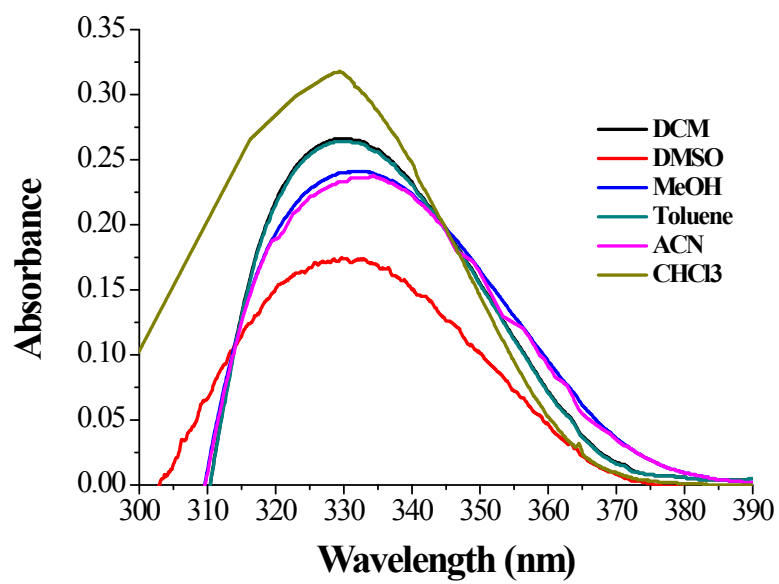
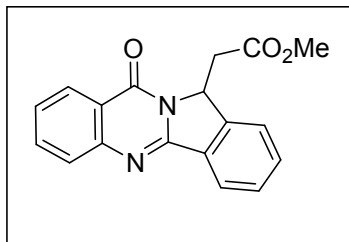


Figure 1: UV-Vis absorption spectra of **3j** in different solvents

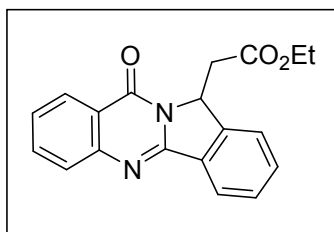
Characterization of data for (3a – 3t):

Methyl 2-(10-oxo-10,12-dihydroisindolo[1,2-b]quinazolin-12-yl)acetate (3a):²



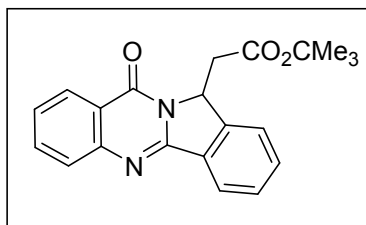
Yellow solid (60.8 mg, 91%), mp. 104 °C. ¹H NMR (400 MHz, Chloroform-*d*): δ 8.36 – 8.32 (m, 1H), 8.16 (d, *J* = 7.0 Hz, 1H), 7.83 – 7.68 (m, 2H), 7.62 – 7.51 (m, 3H), 7.47 (t, *J* = 6.0 Hz, 1H), 5.87 (dd, *J* = 7.8, 3.6 Hz, 1H), 3.72 – 3.67 (m, 1H), 3.66 (s, 3H), 2.98 (dd, *J* = 16.5, 7.9 Hz, 1H). ¹³C NMR (125 MHz, Chloroform-*d*): δ 170.4, 160.8, 154.4, 149.2, 143.3, 134.4, 132.7, 131.9, 129.4, 127.4, 126.6, 126.5, 123.5, 123.2, 121.0, 58.5, 52.0, 35.8. HRMS calcd. for C₁₈H₁₅N₂O₃: (M+H)⁺ 307.1084, found : 307.1083.

Ethyl 2-(10-oxo-10,12-dihydroisindolo[1,2-b]quinazolin-12-yl)acetate (3b):



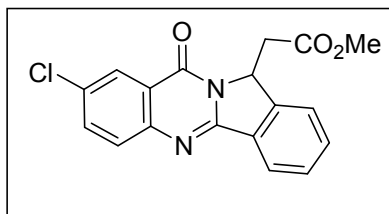
Off white solid (42.4 mg, 85%), mp. 132 °C. ¹H NMR (500 MHz, Chloroform-*d*): δ 8.34 (d, *J* = 7.5 Hz, 1H), 8.16 (d, *J* = 7.5 Hz, 1H), 7.82 – 7.64 (m, 2H), 7.63 – 7.50 (m, 3H), 7.48 (t, *J* = 7.2 Hz, 1H), 5.85 (dd, *J* = 7.7, 3.7 Hz, 1H), 4.08 (q, *J* = 7.2 Hz, 2H), 3.65 (dd, *J* = 16.2, 3.7 Hz, 1H), 3.04 (dd, *J* = 16.2, 7.7 Hz, 1H), 1.12 (t, *J* = 7.2 Hz, 3H). ¹³C NMR (125 MHz, Chloroform-*d*): δ 169.7, 160.7, 154.5, 149.2, 143.3, 134.3, 132.6, 131.9, 129.4, 127.8, 126.6, 126.5, 123.6, 123.3, 121.0, 60.6, 58.6, 35.9, 14.0. HRMS calcd. for C₁₉H₁₇N₂O₃: (M+H)⁺ 321.1241, found : 321.1239.

Tert-butyl 2-(10-oxo-10,12-dihydroisoindolo[1,2-b]quinazolin-12-yl)acetate (3c):



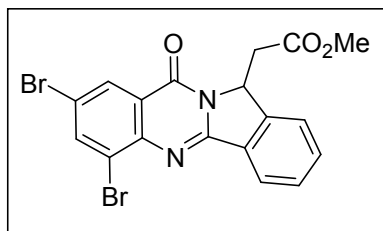
Off white solid (37.7 mg, 77%), mp. 141 °C. ¹H NMR (500 MHz, Chloroform-*d*): δ 8.35 (d, *J* = 7.9 Hz, 1H), 8.17 (d, *J* = 7.4 Hz, 1H), 7.82 – 7.74 (m, 2H), 7.61 – 7.56 (m, 3H), 7.48 (t, *J* = 7.37 Hz, 1H), 5.78 (dd, *J* = 6.9, 3.6 Hz, 1H), 3.46 (dd, *J* = 15.8, 3.6 Hz, 1H), 3.22 – 3.17 (m, 1H), 1.19 (s, 9H). ¹³C NMR (125 MHz, Chloroform-*d*): δ 168.7, 160.6, 154.6, 149.0, 143.2, 134.3, 132.5, 132.0, 129.3, 127.2, 126.5, 126.4, 123.5, 123.3, 121.0, 81.3, 58.8, 36.6, 27.6 (3C). HRMS calcd. for C₂₁H₂₁N₂O₃: (M+H)⁺ 349.1554, found : 349.1552.

Methyl 2-(8-chloro-10-oxo-10,12-dihydroisoindolo[1,2-b]quinazolin-12-yl)acetate (3d):



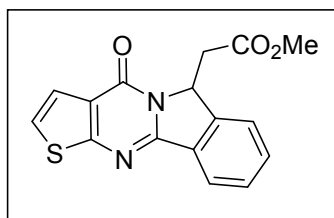
Off white solid (36.2 mg, 72%), mp. 145 °C. ¹H NMR (500 MHz, Chloroform-*d*): δ 8.19 – 8.13 (m, 1H), 8.04 (d, *J* = 6.8 Hz, 1H), 7.75 (d, *J* = 7.5 Hz, 1H), 7.66 – 7.52 (m, 1H), 7.53 – 7.49 (m, 3H), 5.77 – 5.73 (m, 1H), 3.58 (s, 3H), 3.56 (dd, *J* = 14.3, 3.9 Hz, 1H), 2.96 – 2.89 (m, 1H). ¹³C NMR (125 MHz, Chloroform-*d*): δ 170.1, 159.5, 154.6, 147.5, 143.3, 134.7, 132.8, 131.5, 129.5, 128.8, 126.3, 125.9, 123.6, 123.2, 121.9, 58.7, 51.9, 35.5. HRMS calcd. for C₁₈H₁₄ClN₂O₃: (M+H)⁺ 341.0695, found : 341.0694.

Methyl 2-(6,8-dibromo-10-oxo-10,12-dihydroisoindolo[1,2-b]quinazolin-12-yl)acetate (3e):



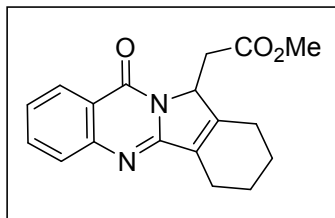
Off white solid (36.1 mg, 70%), mp. 168 °C. ¹H NMR (500 MHz, Chloroform-*d*): δ 8.37 (s, 1H), 8.18 (d, *J* = 7.5 Hz, 1H), 8.11 (s, 1H), 7.63 – 7.54 (m, 2H), 7.45 – 7.42 (m, 1H), 5.81 (dd, *J* = 7.5, 3.5 Hz, 1H), 3.65 (s, 3H), 3.63 – 3.60 (m, 1H), 3.00 (dd, *J* = 16.4, 7.7 Hz, 1H). ¹³C NMR (125 MHz, Chloroform-*d*): δ 170.0, 158.9, 155.2, 146.0, 143.2, 140.3, 133.1, 131.4, 129.5, 128.7, 124.1, 123.5, 123.1, 121.9, 119.3, 58.8, 52.0, 35.3. HRMS calcd. for C₁₈H₁₃Br₂N₂O₃ : (M+H)⁺ 462.9295, found : 462.9294.

Methyl 2-(4-oxo-4,6-dihydrothieno[2',3':4,5]pyrimido[2,1-a]isoindol-6-yl)acetate (3f):



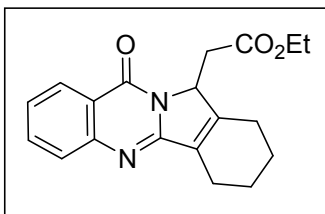
Off white solid (37.2 mg, 78%), mp. 167 °C. ¹H NMR (400 MHz, Chloroform-*d*): δ 8.11 (d, *J* = 6.9 Hz, 1H), 7.81 (d, *J* = 5.3 Hz, 1H), 7.62 – 7.57 (m, 3H), 7.42 (d, *J* = 5.2 Hz, 1H), 5.85 (dd, *J* = 7.9, 3.7 Hz, 1H), 3.73 (dd, *J* = 16.5, 3.7 Hz, 1H), 3.66 (s, 3H), 2.99 (dd, *J* = 16.5, 7.9 Hz, 1H). ¹³C NMR (100 MHz, Chloroform-*d*): δ 170.1, 168.2, 167.1, 155.7, 151.9, 143.4, 141.1, 133.8, 129.9, 128.7, 125.2, 117.4, 116.5, 61.3, 51.6, 46.4. HRMS calcd. for C₁₆H₁₃N₂O₃S: (M+H)⁺ 313.0649, found : 313.0647.

Methyl 2-(10-oxo-1,2,3,4,10,12-hexahydroisindolo[1,2-b]quinazolin-12-yl)acetate (3g):



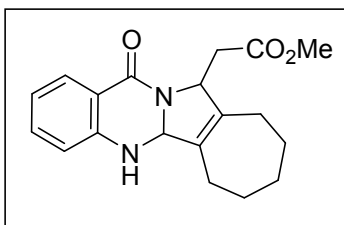
Yellow solid (29.2 mg, 74%), mp. 114 °C. ¹H NMR (400 MHz, Chloroform-*d*): δ 8.29 (d, *J* = 8.1 Hz, 1H), 7.72 – 7.71 (m, 2H), 7.43 – 7.42 (m, 1H), 5.18 (m, 1H), 3.63 (s, 3H), 3.35 (dd, *J* = 15.7, 3.4 Hz, 1H), 3.00 (dd, *J* = 15.1, 7.4 Hz, 1H), 2.51 (m, 2H), 2.36 (m, 2H), 1.85 – 1.74 (m, 2H), 1.32 – 1.24 (m, 2H). ¹³C NMR (100 MHz, Chloroform-*d*): δ 168.9, 160.8, 151.6, 149.3, 134.5, 134.2, 132.7, 127.6, 126.5, 126.0, 121.0, 59.6, 52.2, 38.2, 22.8, 21.9, 21.6, 21.1. HRMS calcd. for C₁₈H₁₉N₂O₃: (M+H)⁺ 311.1397, found : 311.1396.

Ethyl 2-(10-oxo-1,2,3,4,10,12-hexahydroisindolo[1,2-b]quinazolin-12-yl)acetate (3h):



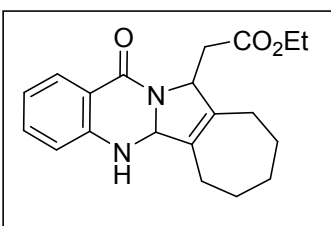
Yellow solid (31.7 mg, 76%), mp. 98 °C. ¹H NMR (400 MHz, Chloroform-*d*): δ 8.29 – 8.22 (m, 1H), 7.71 – 7.69 (m, 2H), 7.43 – 7.39 (m, 1H), 5.15 (m, 1H), 4.08 – 3.98 (m, 2H), 3.26 (dd, *J* = 15.4, 3.8 Hz, 1H), 3.10 (dd, *J* = 15.4, 6.9 Hz, 1H), 2.49 – 2.35 (m, 4H), 1.86 – 1.73 (m, 4H), 1.07 (t, *J* = 7.2 Hz, 3H). ¹³C NMR (100 MHz, Chloroform-*d*): δ 169.6, 160.3, 157.9, 152.3, 149.4, 134.5, 134.1, 131.8, 127.2, 126.5, 126.0, 61.3, 60.9, 33.6, 23.5, 22.1, 21.6, 20.4, 14.1. HRMS calcd. for C₁₉H₂₁N₂O₃: (M+H)⁺ 325.1554, found : 325.1552.

Methyl 2-(13-oxo-5,5a,7,8,9,10,11,13-octahydro-6H-cyclohepta[3,4]pyrrolo[2,1-b]quinazolin-11-yl)acetate (3i):



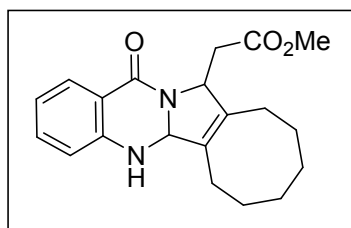
Yellow solid (41.9 mg, 83%), mp. 133 °C. ¹H NMR (400 MHz, Chloroform-*d*): δ 7.99 (d, *J* = 7.5 Hz, 1H), 7.38 (t, *J* = 7.4 Hz, 1H), 6.92 (t, *J* = 7.5 Hz, 1H), 6.59 (d, *J* = 8.3 Hz, 1H), 5.94 (s, 1H), 5.30 (s, 1H), 4.47 (d, *J* = 6.2 Hz, 1H), 3.70 (s, 3H), 2.92 – 2.87 (m, 1H), 2.50 (d, *J* = 17.0 Hz, 1H), 2.26 – 2.09 (m, 2H), 1.78 – 1.77 (m, 2H), 1.51 – 1.32 (m, 4H), 1.27 – 1.24 (m, 2H). ¹³C NMR (100 MHz, Chloroform-*d*): δ 172.4, 166.0, 148.2, 135.9, 133.9, 129.3, 129.1, 119.4, 118.1, 111.4, 67.1, 53.6, 52.3, 35.1, 33.8, 32.1, 31.0, 26.8, 25.4. HRMS calcd. for C₁₉H₂₃N₂O₃ : (M+H)⁺ 327.1710, found : 327.1711.

Ethyl 2-(13-oxo-5,5a,7,8,9,10,11,13-octahydro-6H-cyclohepta[3,4]pyrrolo[2,1-b]quinazolin-11-yl)acetate (3j):



Yellow solid (39.7 mg, 80%), mp. 130 °C. ¹H NMR (400 MHz, DMSO-*d*⁶): δ 8.09 (s, 1H), 7.75 (d, *J* = 6.9 Hz, 1H), 7.38 (t, *J* = 7.8 Hz, 1H), 6.86 (t, *J* = 7.4 Hz, 1H), 6.80 (d, *J* = 8.3 Hz, 1H), 5.05 (s, 1H), 4.75 – 4.74 (m, 1H), 4.10 (q, *J* = 7.1 Hz, 2H), 2.79 – 2.76 (m, 1H), 2.46 (m, 1H), 2.42 – 2.25 (m, 3H), 2.04 – 1.98 (m, 1H), 1.94 – 1.88 (m, 1H), 1.81 – 1.78 (m, 1H), 1.61 – 1.55 (m, 4H), 1.15 (t, *J* = 7.0 Hz, 3H). ¹³C NMR (100 MHz, Chloroform-*d*): δ 171.9, 166.0, 148.3, 135.8, 133.8, 129.3, 129.1, 119.3, 118.1, 111.4, 67.1, 61.4, 53.7, 35.2, 33.8, 32.1, 31.0, 26.8, 25.4, 14.4. HRMS calcd. for C₂₀H₂₅N₂O₃ : (M+H)⁺ 341.1867, found : 341.1866.

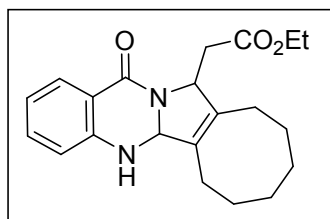
Methyl 2-(14-oxo-5,5a,6,7,8,9,10,11,12,14-decahydrocycloocta[3,4]pyrrolo[2,1-b]quinazolin-12-yl)acetate (3k):



White solid (32.6 mg, 73%), mp. 159 °C. ¹H NMR (400 MHz, DMSO-*d*⁶): δ 8.24 (s, 1H), 7.75 (d, *J* = 7.5 Hz, 1H), 7.37 (t, *J* = 7.8 Hz, 1H), 6.87 (t, *J* = 7.4 Hz, 1H), 6.80 (d, *J* = 8.3 Hz, 1H), 5.11 (s, 1H), 4.79 (d, *J* = 5.2 Hz, 1H), 3.61 (s, 3H), 2.67 (m, 1H), 2.50 (s, 1H), 2.39 – 2.31 (m, 2H), 2.16 (d, *J* = 14.4 Hz, 1H), 1.87 (d, *J* = 12.2 Hz, 1H), 1.64 – 1.53 (m, 2H), 1.34 (m, 4H), 1.23 – 1.21 (m, 2H). ¹³C NMR (125 MHz, Chloroform-*d*): δ 172.4, 165.8, 148.3, 133.6, 133.4, 129.2, 126.3, 119.2, 118.2, 111.3, 64.9, 53.8, 52.2, 31.4, 31.3, 29.3, 27.8, 26.2, 25.9, 25.8.

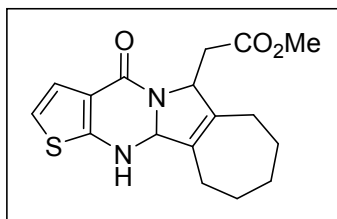
HRMS calcd. for C₂₀H₂₅N₂O₃ : (M+H)⁺ 341.1867, found : 341.1865.

Ethyl 2-(14-oxo-5,5a,6,7,8,9,10,11,12,14-decahydrocycloocta[3,4]pyrrolo[2,1-b]quinazolin-12-yl)acetate (3l):



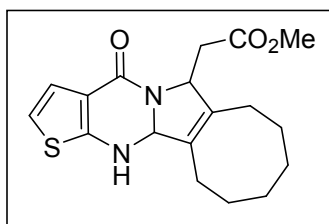
Yellow solid (37.7 mg, 72%), mp. 123 °C. ¹H NMR (400 MHz, Chloroform-*d*): δ 7.97 (dd, *J* = 7.8, 1.6 Hz, 1H), 7.37 (ddd, *J* = 8.3, 7.3, 1.7 Hz, 1H), 6.90 (t, *J* = 8.0 Hz, 1H), 6.58 (d, *J* = 8.0 Hz, 1H), 5.35 (s, 1H), 4.45 (d, *J* = 8.2 Hz, 1H), 4.19 – 4.10 (m, 2H), 2.61 (t, *J* = 17.1 Hz, 2H), 2.41 (t, *J* = 9.9 Hz, 2H), 2.12 (d, *J* = 14.4 Hz, 1H), 1.48 – 1.36 (m, 9H), 1.22 (t, *J* = 7.1 Hz, 3H). ¹³C NMR (100 MHz, Chloroform-*d*): δ 172.0, 166.1, 148.5, 133.9, 133.5, 129.3, 126.2, 119.3, 118.1, 111.5, 65.0, 61.5, 53.9, 31.5, 31.3, 29.4, 27.9, 26.4, 26.0 (2C), 14.3. HRMS calcd. for C₂₁H₂₇N₂O₃ : (M+H)⁺ 355.2023, found : 355.2022.

Methyl 2-(4-oxo-6,7,8,9,10,11,11b,12-octahydro-4H-cyclohepta[3,4]pyrrolo[1,2-a]thieno[2,3-d]pyrimidin-6-yl)acetate (3m):



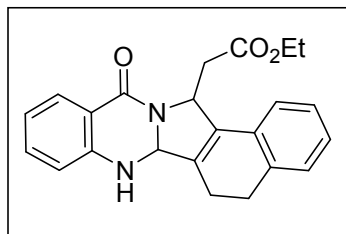
Yellow solid (34.1 mg, 70%), mp. 148 °C. ¹H NMR (400 MHz, DMSO-*d*⁶): δ 7.72 (d, *J* = 5.3 Hz, 1H), 7.56 (s, 1H), 6.89 (d, *J* = 5.3 Hz, 1H), 5.01 (s, 1H), 4.78 (d, *J* = 5.4 Hz, 1H), 3.62 (s, 3H), 2.73 (m, 1H), 2.49 – 2.22 (m, 3H), 2.04 – 1.91 (m, 2H), 1.78 (m, 1H), 1.61 – 1.59 (m, 3H), 1.20 – 1.13 (m, 2H). ¹³C NMR (100 MHz, Chloroform-*d*): δ 172.0, 163.6, 154.0, 136.2, 133.7, 129.0, 114.7, 110.2, 68.5, 55.5, 52.5, 35.2, 33.5, 32.1, 29.2, 26.8, 25.4. HRMS calcd. for C₁₇H₂₁N₂O₃S: (M+H)⁺ 333.1275, found : 333.1276.

Methyl 2-(4-oxo-4,6,7,8,9,10,11,12,12b,13-decahydrocycloocta[3,4]pyrrolo[1,2-a]thieno[2,3-d]pyrimidin-6-yl)acetate (3n):



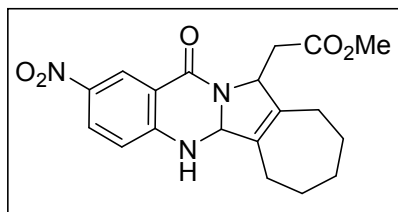
Off white solid (36.2 mg, 73%), mp. 139 °C. ¹H NMR (400 MHz, Chloroform-*d*): δ 7.48 (d, *J* = 5.3 Hz, 1H), 6.53 (d, *J* = 5.2 Hz, 1H), 5.40 (s, 1H), 5.34 (s, 1H), 4.41 (d, *J* = 5.0 Hz, 1H), 3.70 (s, 3H), 2.79 (dd, *J* = 16.8, 5.9 Hz, 1H), 2.58 (d, *J* = 16.8 Hz, 1H), 2.41 – 2.37 (m, 2H), 2.15 – 2.12 (m, 1H), 2.01 – 1.97 (m, 1H), 1.70 – 1.60 (m, 2H), 1.56 – 1.37 (m, 6H). ¹³C NMR (100 MHz, Chloroform-*d*): δ 172.0, 163.9, 154.7, 139.3, 134.6, 133.9, 125.8, 114.5, 66.6, 55.9, 52.5, 31.7, 31.2, 31.0, 27.8, 26.4, 26.2, 25.9. HRMS calcd. for C₁₈H₂₃N₂O₃S: (M+H)⁺ 347.1431, found : 347.1430.

Ethyl 2-(12-oxo-5,6,6b,7,12,14-hexahydrobenzo[4,5]isoindolo[1,2-b]quinazolin-14-yl)acetate (3o):



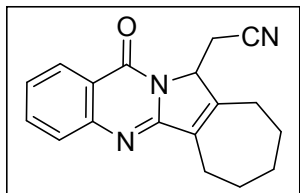
Yellow solid (22.8 mg, 55%), mp. 145 °C. ¹H NMR (400 MHz, Chloroform-*d*): δ 8.02 (d, *J* = 7.8 Hz, 1H), 7.43 (t, *J* = 7.6 Hz, 1H), 7.37 – 7.17 (m, 3H), 6.97 (t, *J* = 7.4 Hz, 1H), 6.68 (d, *J* = 8.0 Hz, 1H), 6.04 (s, 1H), 5.69 (s, 1H), 4.71 – 4.70 (m, 1H), 4.20 – 4.08 (m, 2H), 3.15 – 3.00 (m, 2H), 2.85 – 2.83 (m, 2H), 2.52 – 2.39 (m, 1H), 2.20 – 2.16 (m, 1H), 1.18 (t, *J* = 7.1 Hz, 3H). ¹³C NMR (125 MHz, Chloroform-*d*): δ 171.5, 168.1, 160.8, 149.3, 137.7, 134.5, 134.0, 130.9, 129.9, 128.5, 127.6, 127.1, 126.7, 126.4, 124.7, 122.5, 66.3, 60.9, 53.3, 29.7, 27.8, 18.2, 13.7. HRMS calcd. for C₂₃H₂₃N₂O₃ : (M+H)⁺ 375.1710, found : 375.1711.

Methyl 2-(2-nitro-13-oxo-5,5a,7,8,9,10,11,13-octahydro-6H-cyclohepta[3,4]pyrrolo[2,1-b]quinazolin-11-yl)acetate (3p)^a:



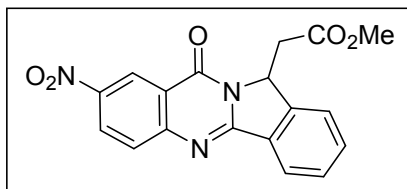
Yellow solid (25 mg, 62%), mp. 168 °C. ¹H NMR (400 MHz, DMSO-*d*⁶) δ 8.50 (d, *J* = 2.8 Hz, 2H), 8.17 (dd, *J* = 9.3, 2.8 Hz, 1H), 7.10 (d, *J* = 9.6 Hz, 1H), 5.10 (s, 1H), 5.03 (d, *J* = 5.6 Hz, 1H), 3.61 (s, 3H), 2.79 – 2.74 (m, 1H), 2.38 – 2.23 (m, 3H), 2.04 – 1.96 (m, 1H), 1.87 (dd, *J* = 15.1, 6.5 Hz, 1H), 1.78 – 1.74 (m, 2H), 1.60 – 1.50 (m, 4H). ¹³C NMR (75 MHz, DMSO-*d*⁶) δ 170.4, 169.3, 162.7, 155.4, 152.4, 129.3, 126.1, 123.9, 116.9, 113.1, 66.2, 53.8, 52.1, 34.1, 32.4, 31.0, 28.8, 26.0, 24.0. HRMS calcd. for C₁₉H₂₂N₃O₅ : (M+H)⁺ 372.1561, found : 372.1562.

2-(13-oxo-7,8,9,10,11,13-hexahydro-6H-cyclohepta[3,4]pyrrolo[2,1-b]quinazolin-11-yl)acetonitrile (3q):



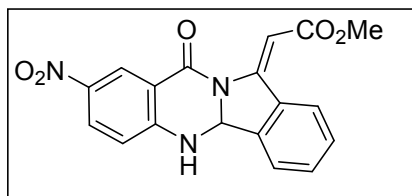
Yellow solid (22 mg, 74%), mp. 134 °C. ¹H NMR (400 MHz, Chloroform-*d*) δ 7.83 (dd, *J* = 7.7, 1.3 Hz, 1H), 7.25 – 7.23 (m, 1H), 6.78 (t, *J* = 7.3 Hz, 1H), 6.60 (d, *J* = 8.0 Hz, 2H), 2.04 – 1.99 (m, 4H), 1.60 – 1.58 (m, 3H), 1.55 – 1.49 (m, 3H), 1.27 – 1.23 (m, 2H). ¹³C NMR (101 MHz, Chloroform-*d*) δ 164.2, 145.7, 134.0, 132.0, 131.0, 128.9, 128.4, 118.8, 117.7, 115.2, 114.9, 72.6, 41.6 (2C), 29.1 (2C), 21.6 (2C). HRMS calcd. for C₁₈H₁₈N₃O : (M+H)⁺ 292.1452, found : 292.1450.

Methyl 2-(8-nitro-10-oxo-10,12-dihydroisindolo[1,2-b]quinazolin-12-yl)acetate (3r)^a:



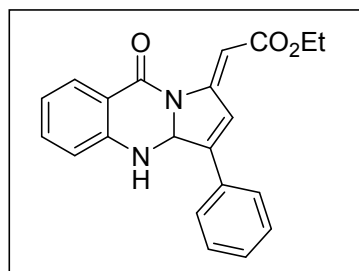
Brown liquid (16 mg, 32%). ¹H NMR (400 MHz, Chloroform-*d*) δ 9.19 (d, *J* = 2.6 Hz, 1H), 8.55 (dd, *J* = 9.3, 2.5 Hz, 1H), 8.22 (d, *J* = 7.7 Hz, 1H), 7.93 (d, *J* = 9.0 Hz, 1H), 7.70 – 7.63 (m, 3H), 5.87 (dd, *J* = 7.4, 3.6 Hz, 1H), 3.63 (s, 3H), 3.61 – 3.59 (m, 1H), 3.14 – 3.08 (m, 1H). ¹³C NMR (101 MHz, Chloroform-*d*) δ 170.0, 159.6, 157.6, 153.3, 145.4, 143.8, 134.0, 129.9, 128.7, 128.6, 124.4, 123.4, 123.3, 121.6, 121.4, 59.2, 52.2, 35.3. HRMS calcd. for C₁₈H₁₄N₃O₅ : (M+H)⁺ 352.0935, found : 352.0937.

Methyl (E)-2-(8-nitro-10-oxo-5,10-dihydroisoindolo[1,2-b]quinazolin-12(4bH)-ylidene)acetate (3s)^a:



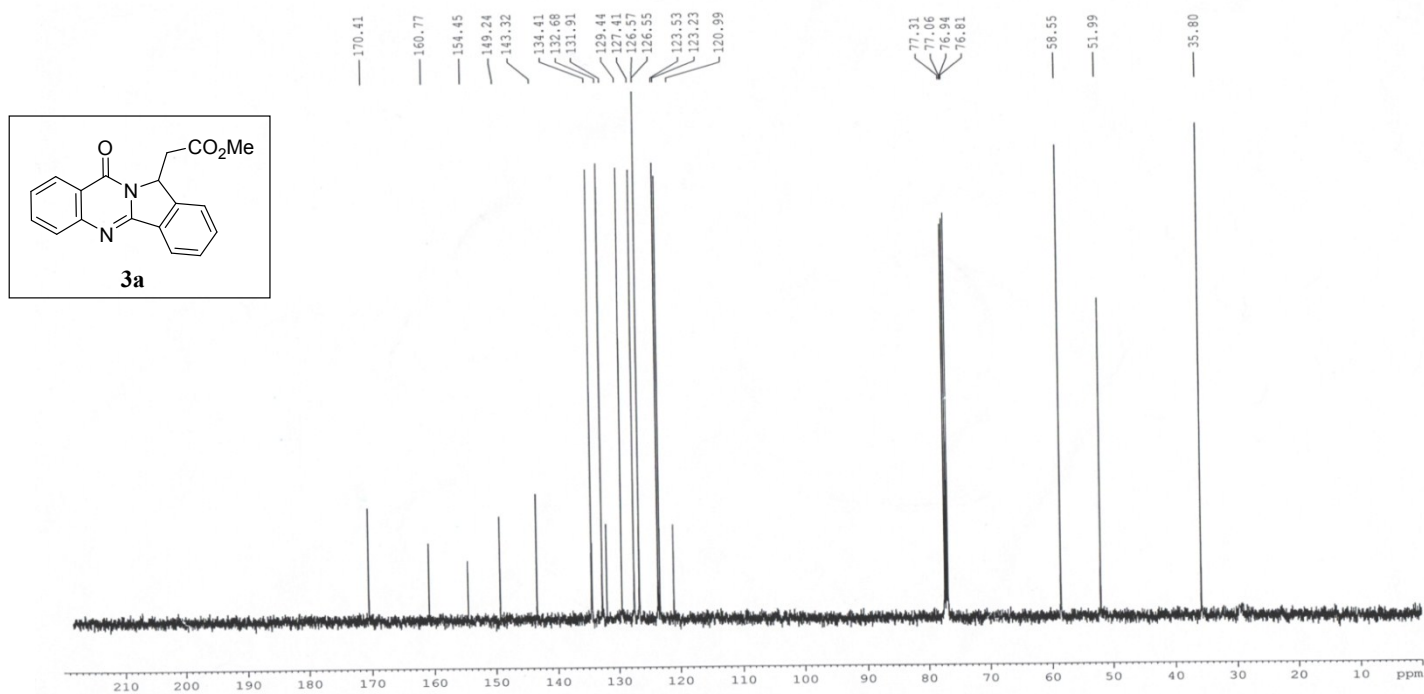
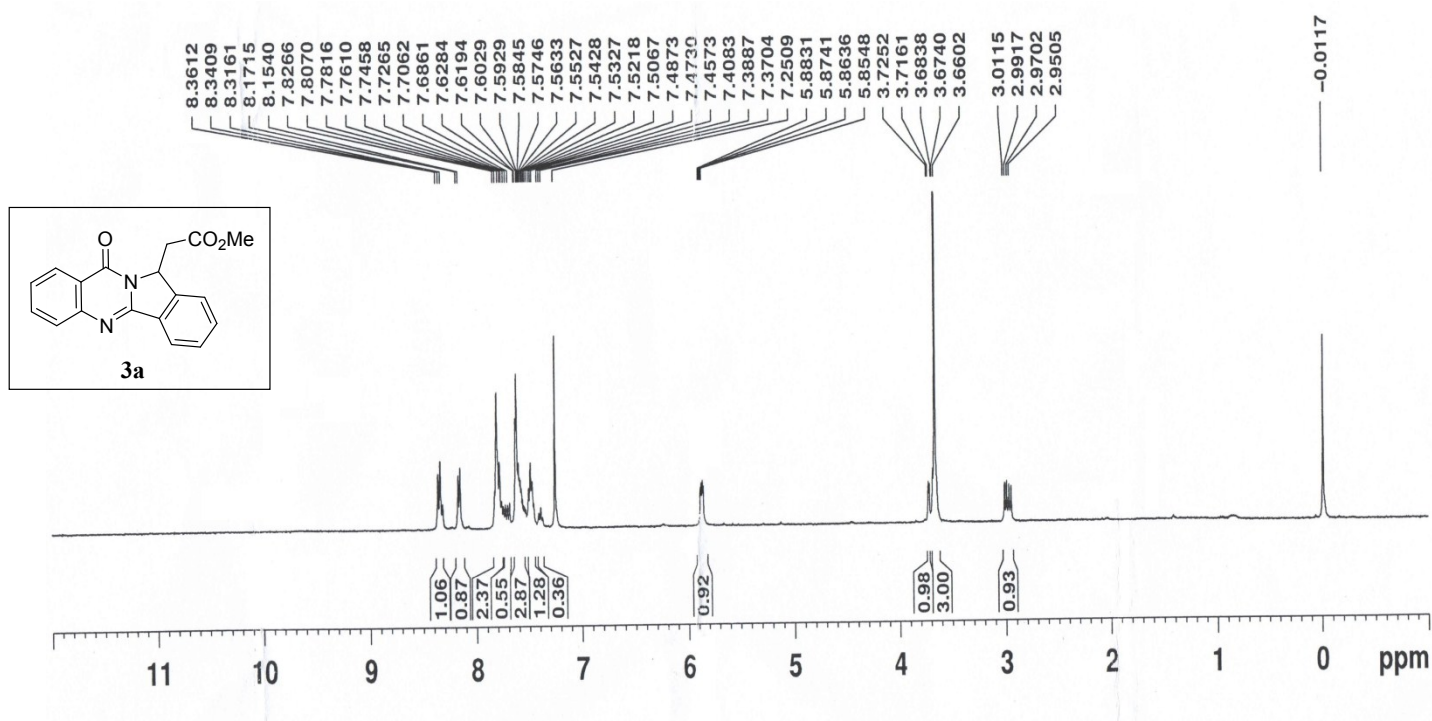
Off white solid (35 mg, 53%), mp. 152 °C. ¹H NMR (500 MHz, 12% DMSO-*d*⁶ in Chloroform-*d*) δ 8.61 (d, *J* = 2.5 Hz, 1H), 8.12 (d, *J* = 15.6 Hz, 1H), 8.00 (dd, *J* = 9.0, 2.6 Hz, 1H), 7.77 (s, 1H), 7.64 (d, *J* = 7.7 Hz, 1H), 7.55 (d, *J* = 7.0 Hz, 1H), 7.43 – 7.37 (m, 2H), 6.73 (d, *J* = 9.0 Hz, 1H), 6.28 (d, *J* = 4.9 Hz, 1H), 3.67 (s, 3H). ¹³C NMR (126 MHz, 12% DMSO-*d*⁶ in Chloroform-*d*) δ 166.0, 162.1, 152.0, 140.4, 138.0, 137.2, 133.1, 129.8, 129.3, 128.5, 128.0, 126.9, 124.8, 121.0, 114.0, 112.4, 64.6, 51.2. HRMS calcd. for C₁₈H₁₄N₃O₅ : (M+H)⁺ 352.0935, found : 352.0934.

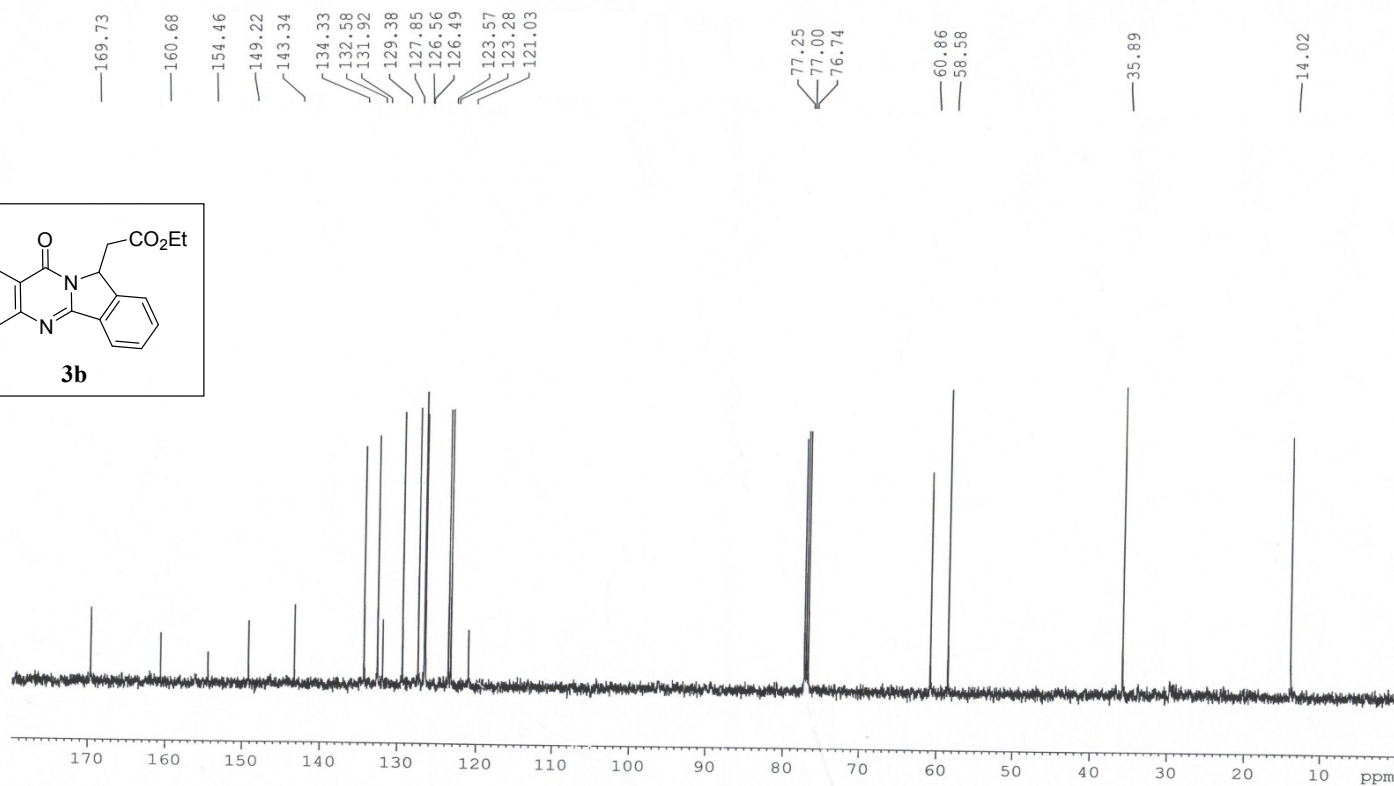
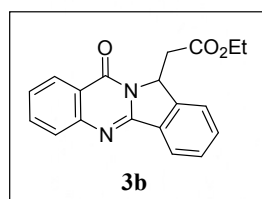
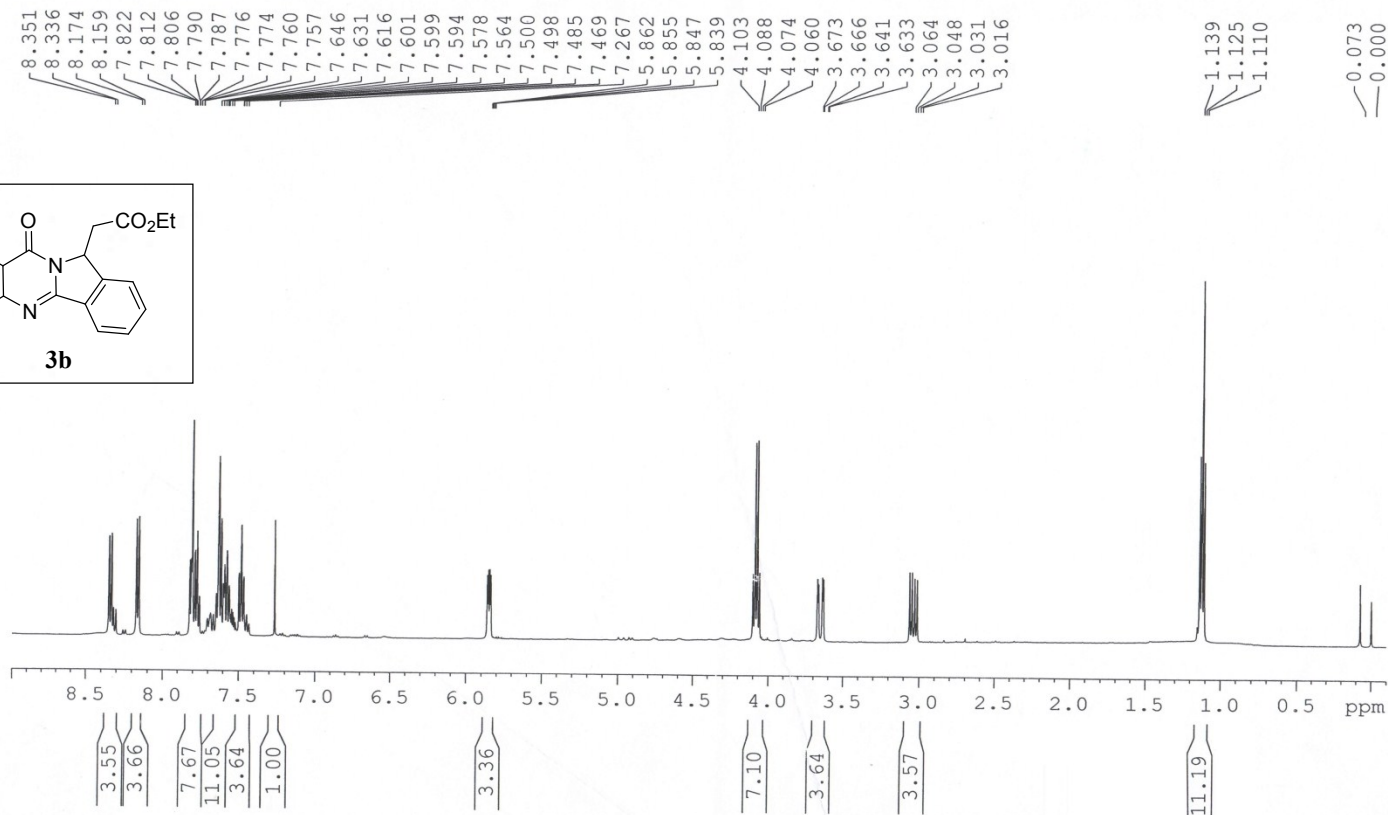
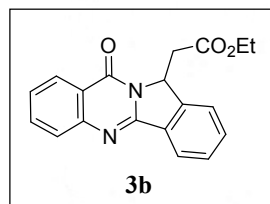
Ethyl (E)-2-(9-oxo-3-phenyl-4,9-dihydropyrrolo[2,1-b]quinazolin-1(3aH)-ylidene)acetate (3t):

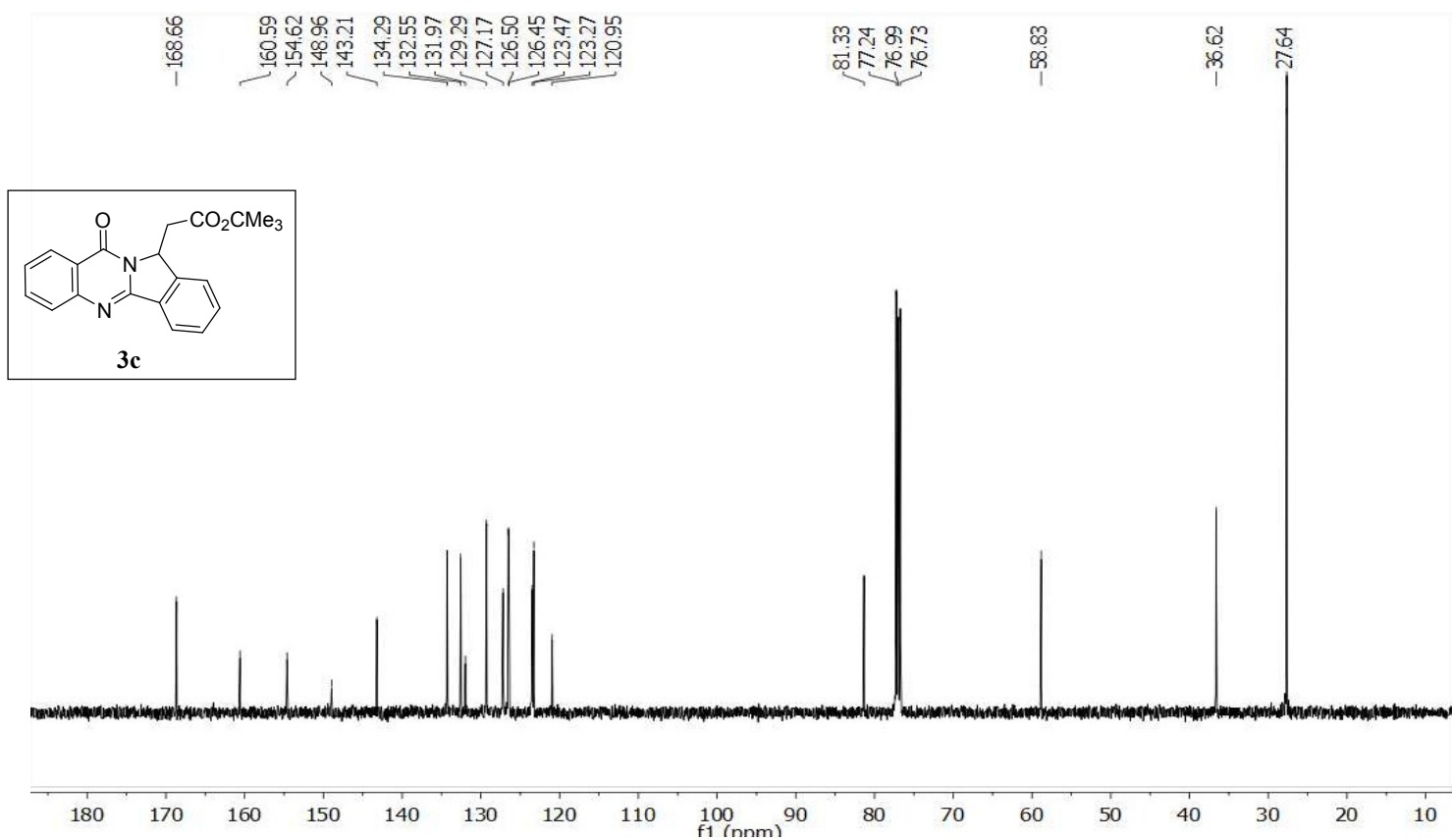
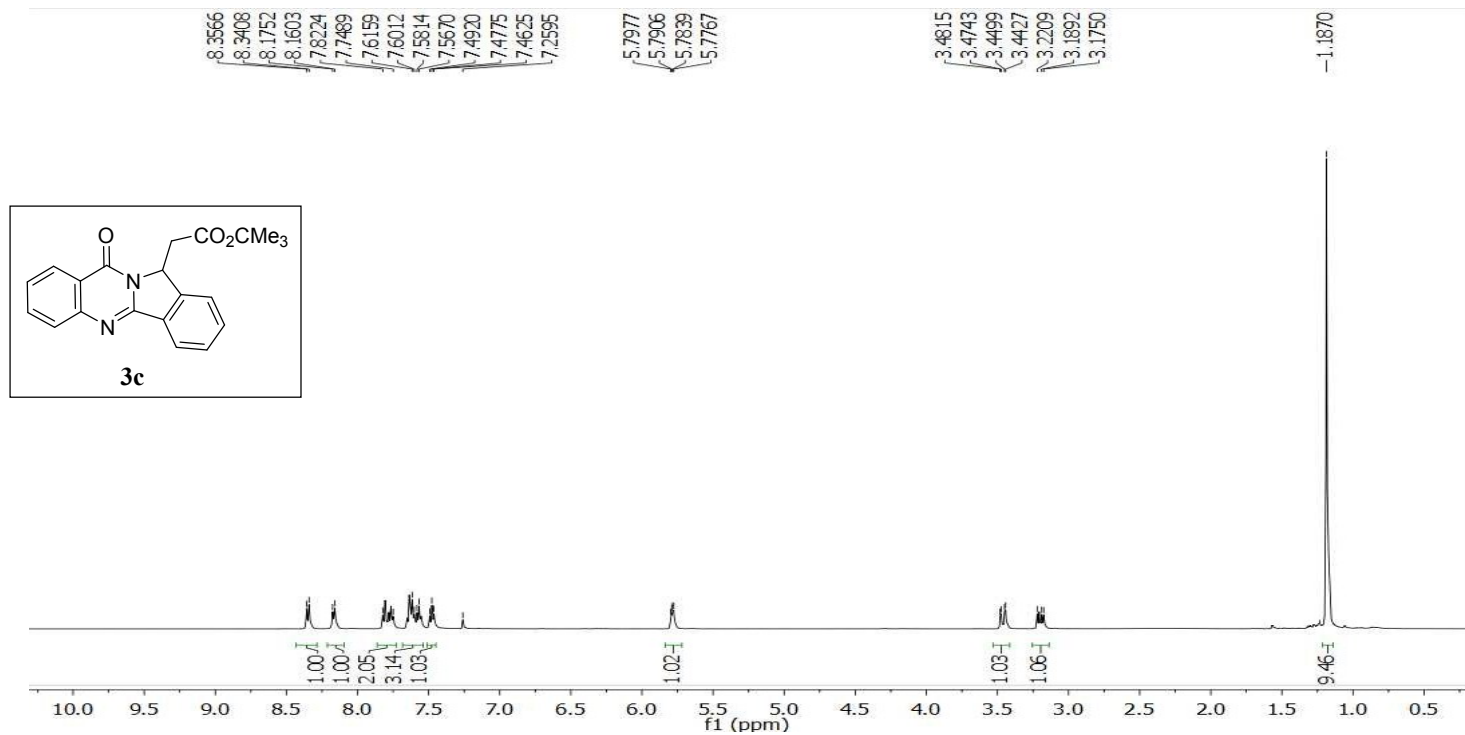


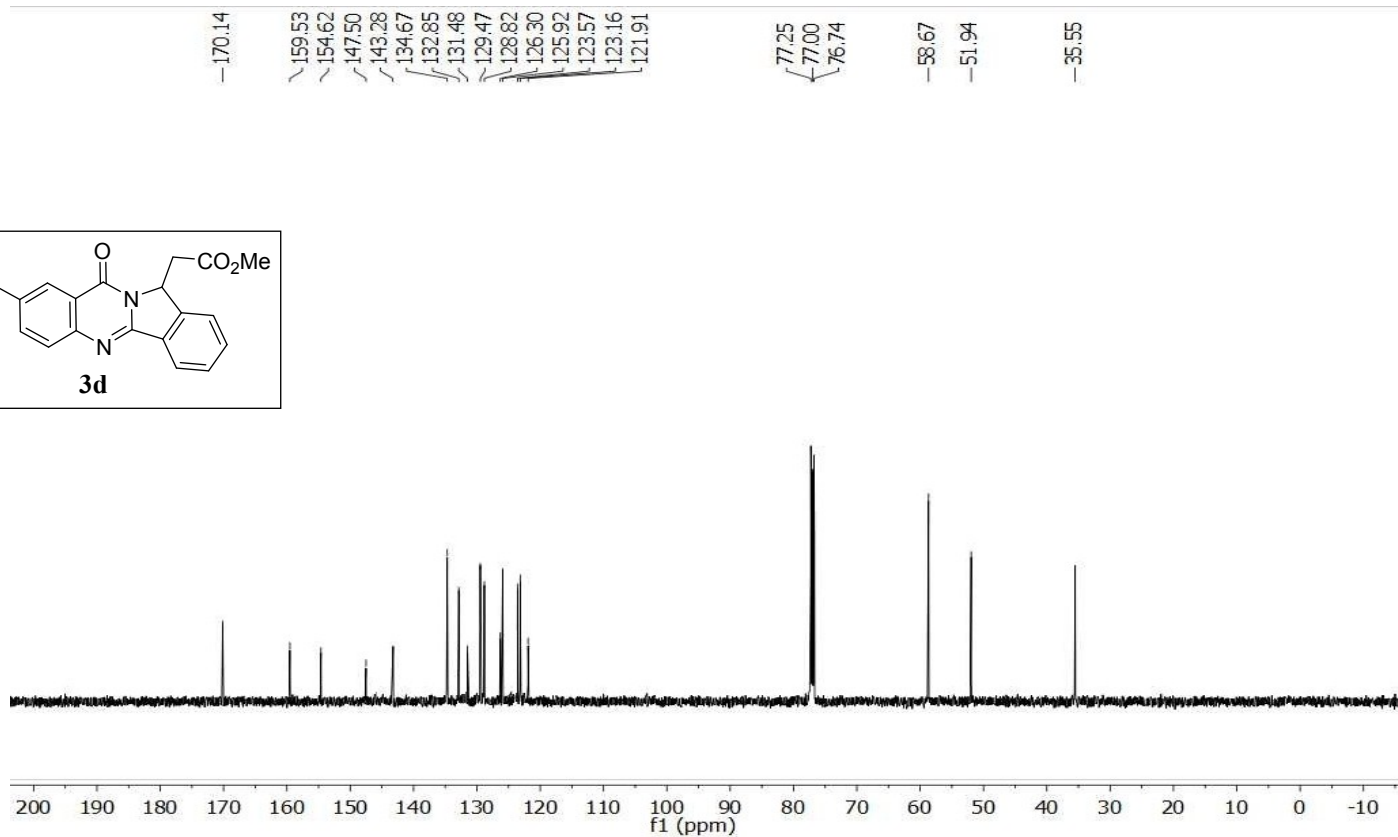
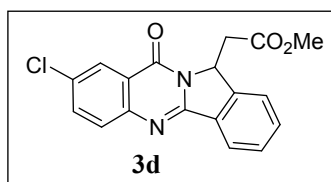
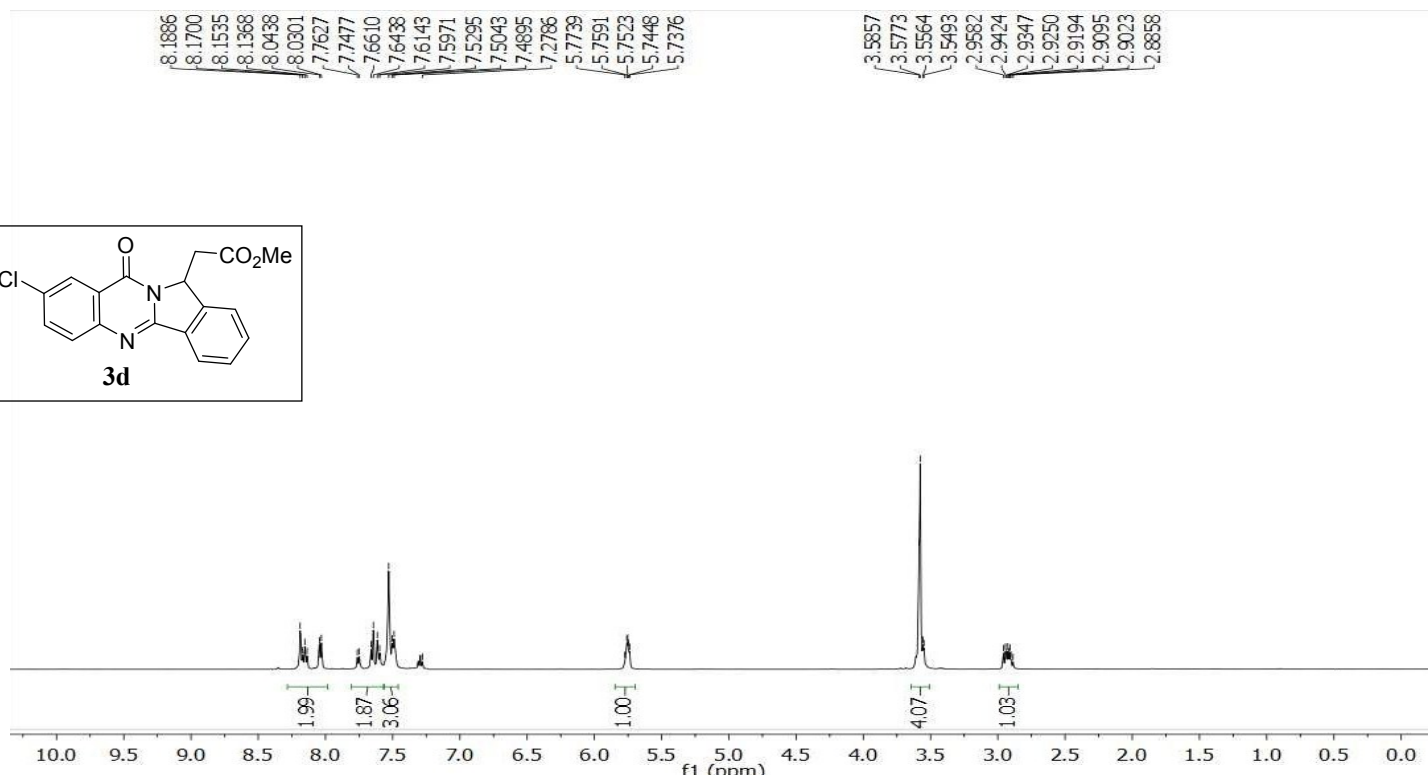
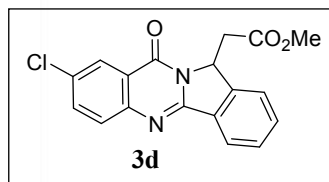
Yellow solid (35.8 mg, 72%), mp. 72 °C. ¹H NMR (400 MHz, DMSO-*d*⁶): δ 8.15 (s, 1H), 7.58 – 7.54 (m, 1H), 7.50 – 7.39 (m, 2H), 7.29 (d, *J* = 6.9 Hz, 2H), 7.24 (t, *J* = 7.1 Hz, 1H), 6.80 (s, 1H), 6.71 – 6.66 (m, 2H), 6.47 (d, *J* = 9.4 Hz, 1H), 5.33 (d, *J* = 15.6 Hz, 1H), 4.89 (d, *J* = 9.5 Hz, 1H), 4.08 (q, *J* = 7.2 Hz, 2H), 1.17 (td, *J* = 7.0, 3.2 Hz, 3H). ¹³C NMR (100 MHz, Chloroform-*d*): δ 166.6, 164.8, 146.6, 146.4, 143.3, 135.3, 134.5, 134.0, 129.0 (2C), 128.9 (2C), 128.7, 128.6, 123.9, 119.5, 115.5, 114.9, 63.2, 60.7, 14.3. HRMS calcd. for C₂₁H₁₉N₂O₃ : (M+H)⁺ 347.1397, found : 347.1396.

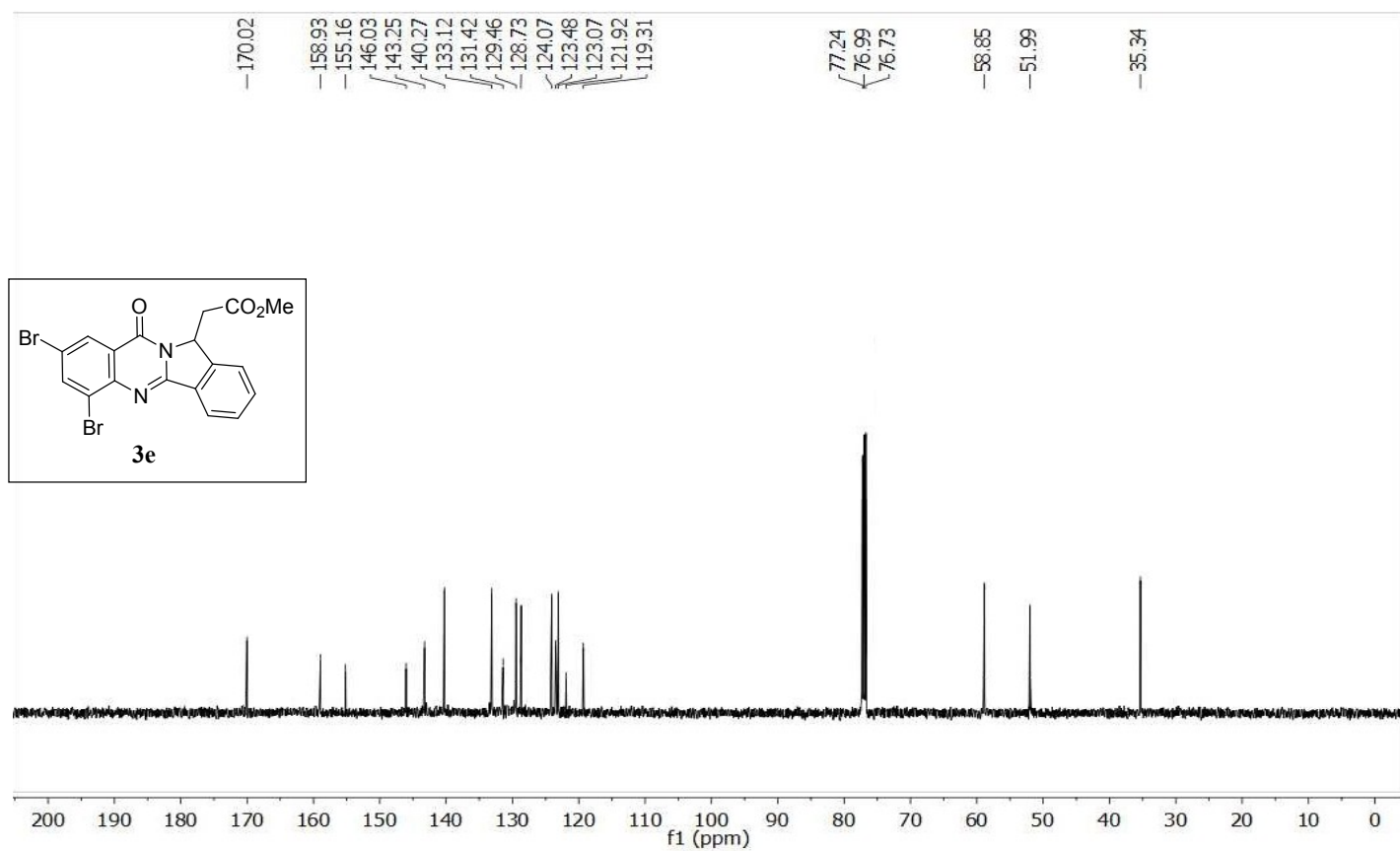
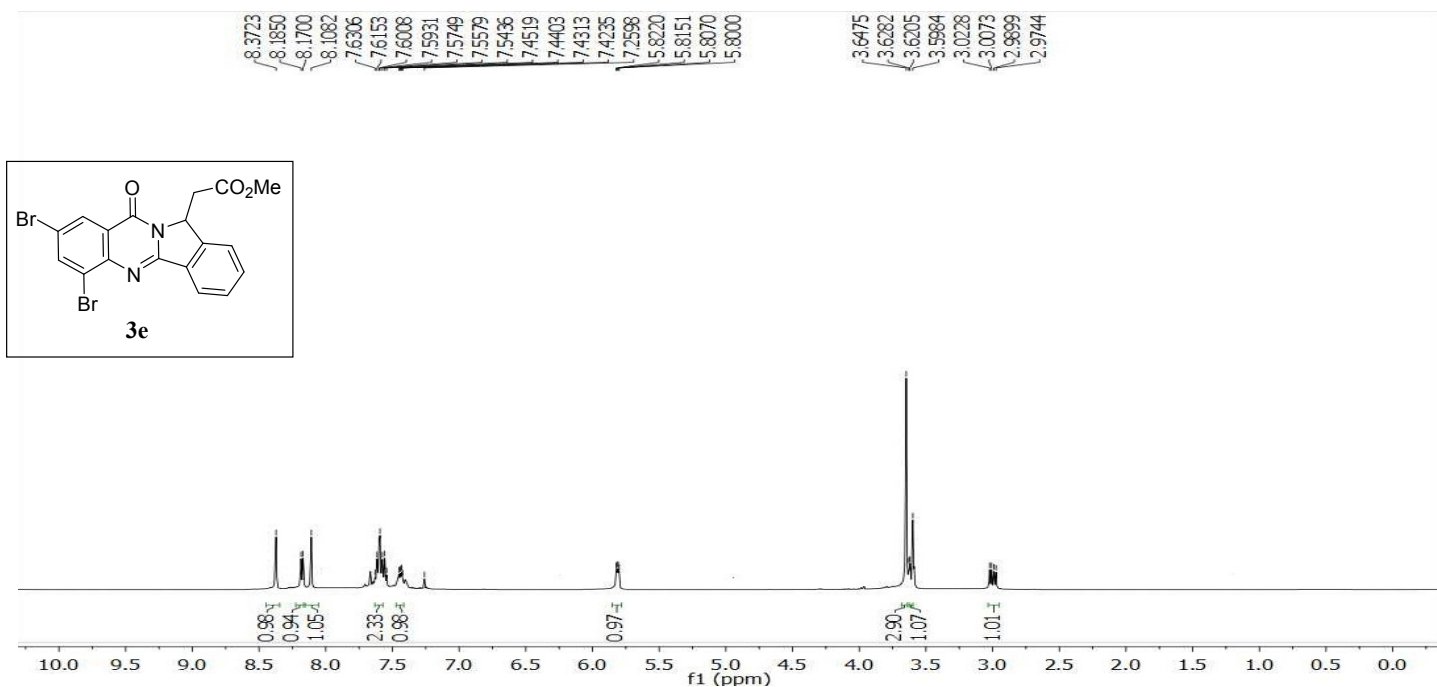
¹H and ¹³C Spectra of (3a – 3t):

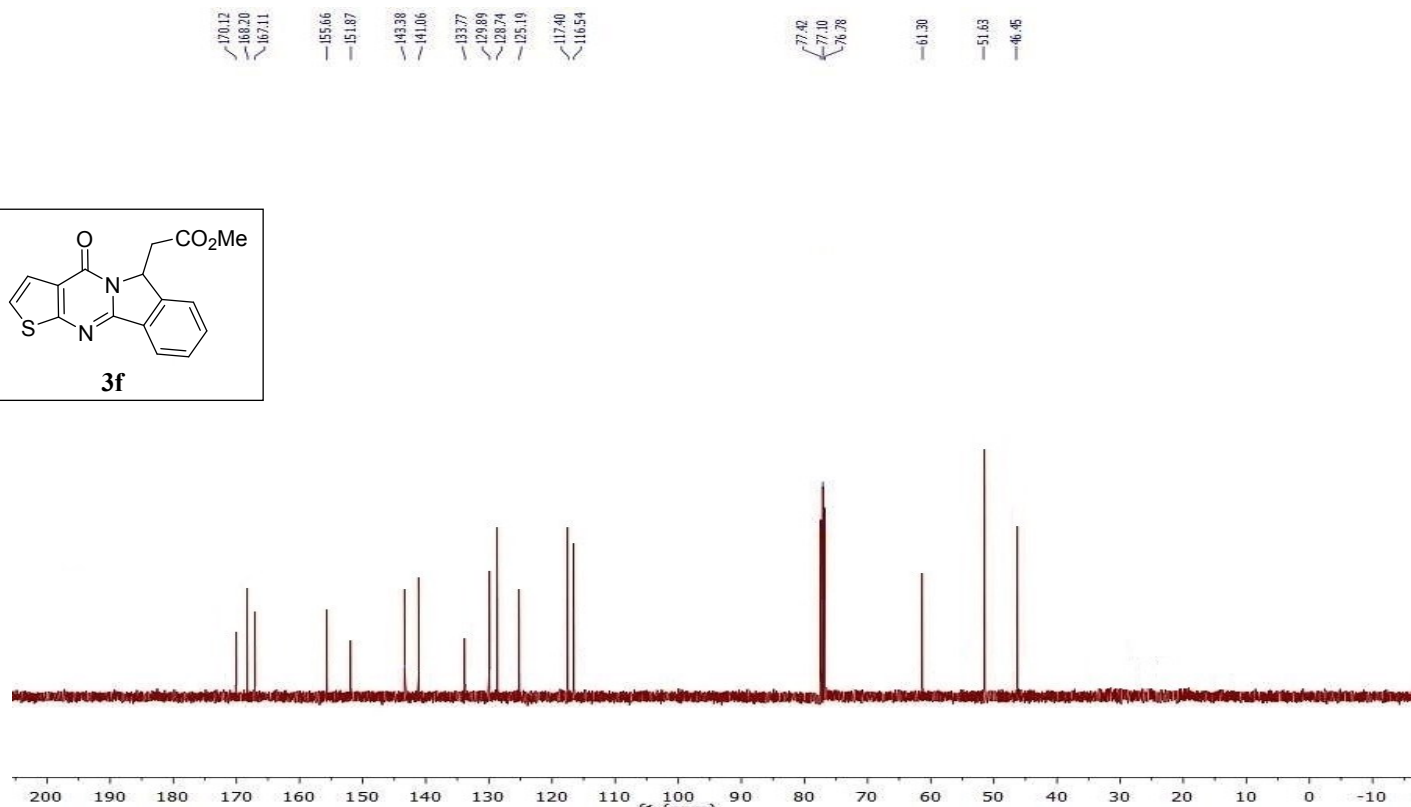
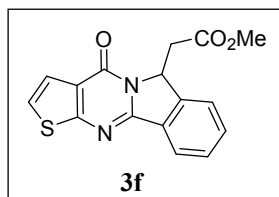
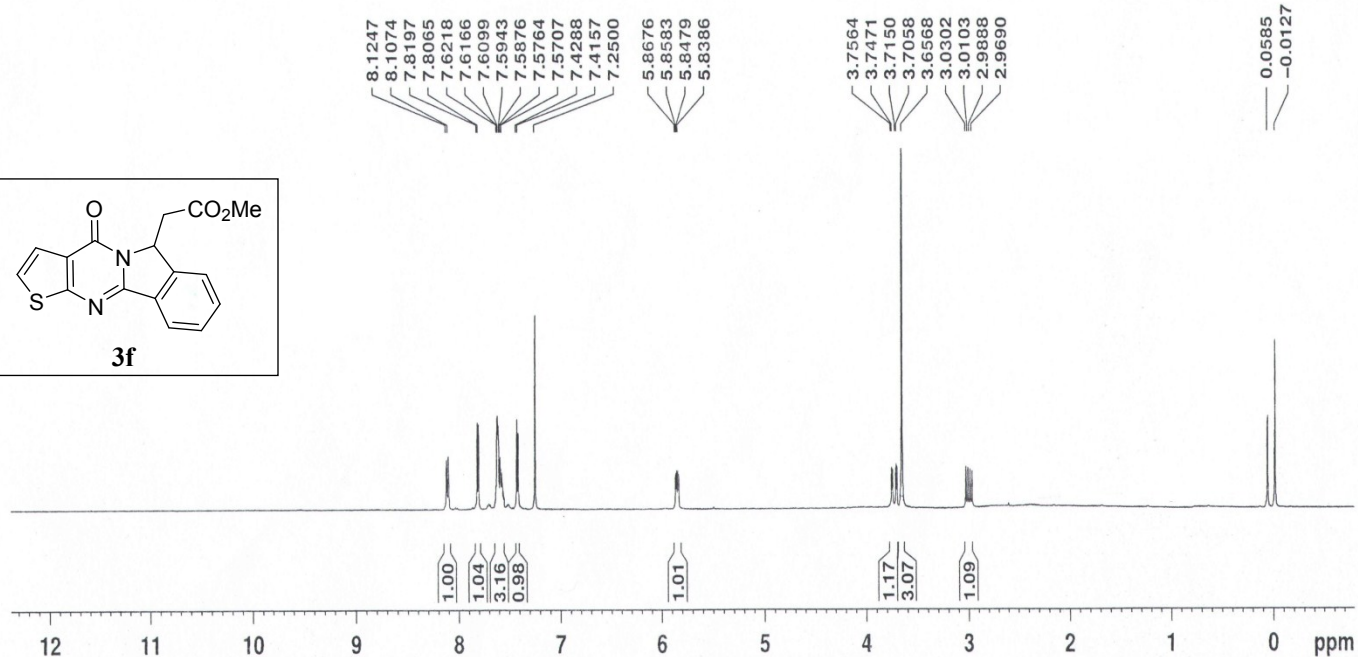
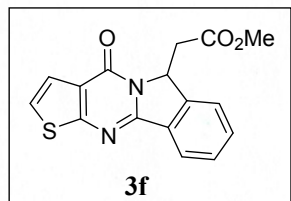


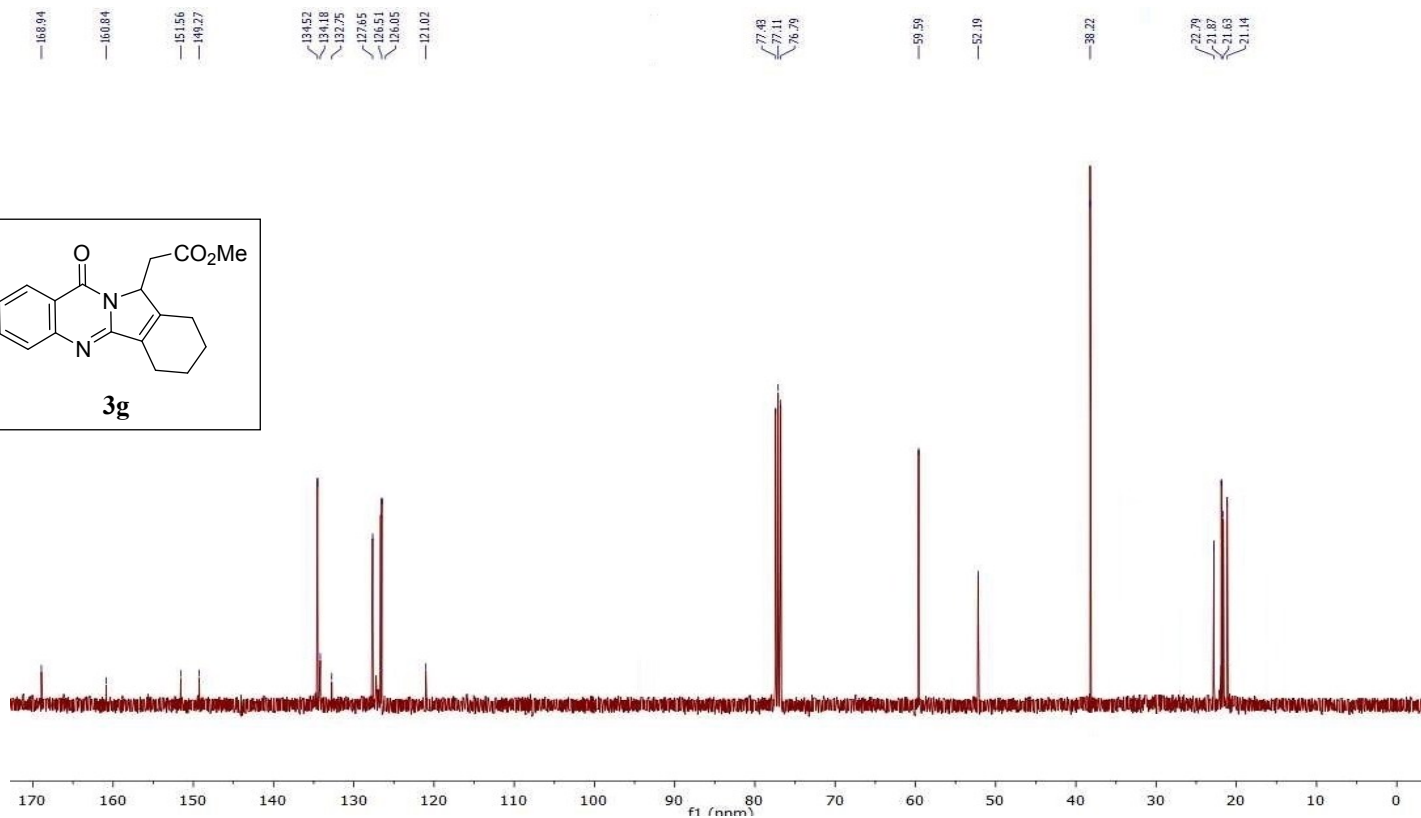
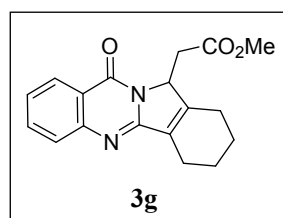
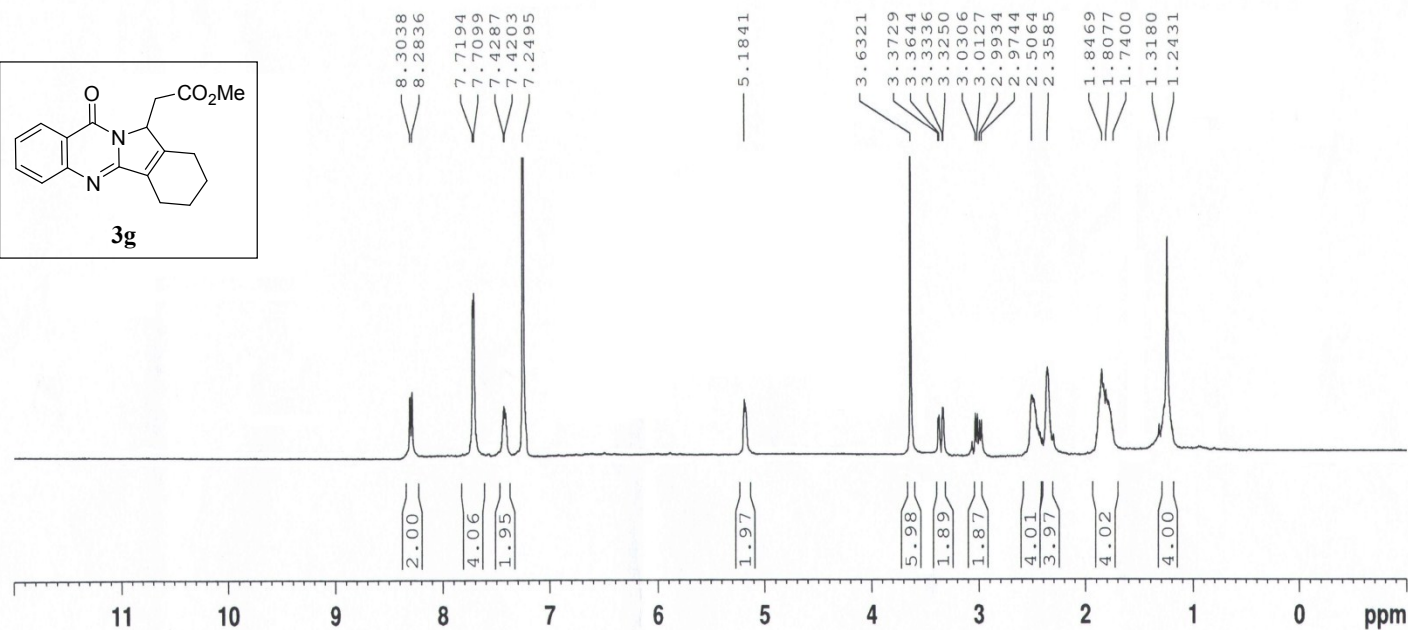
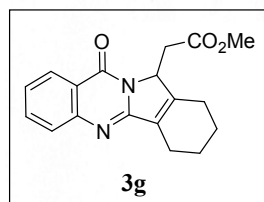


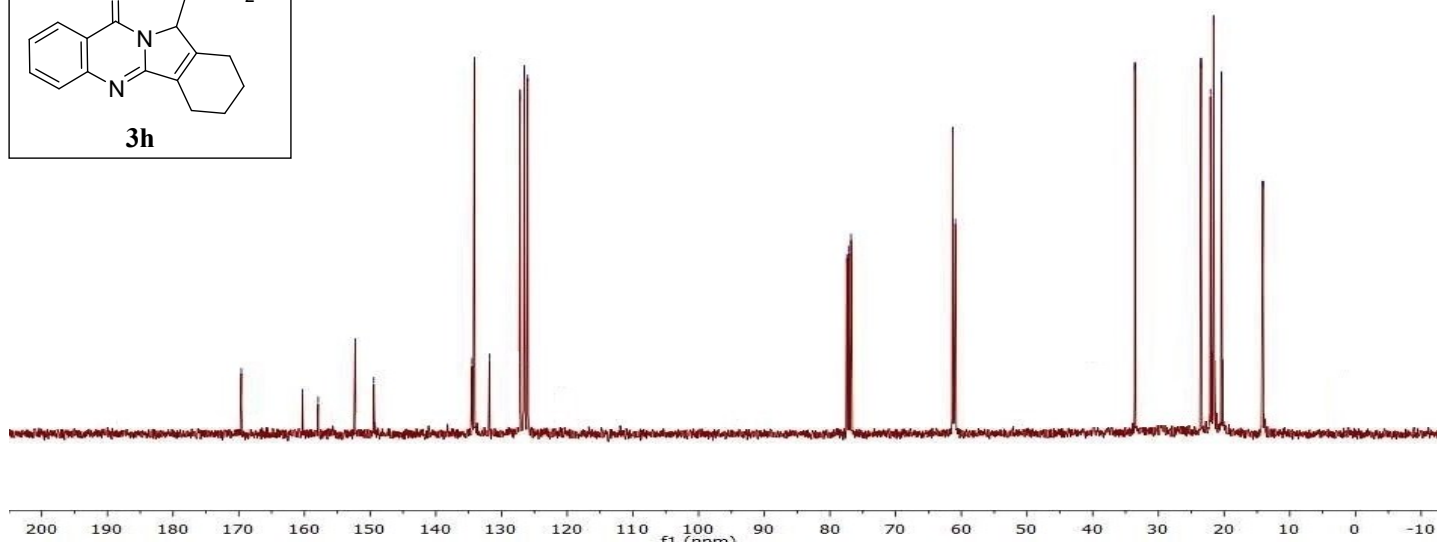
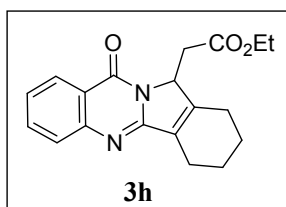
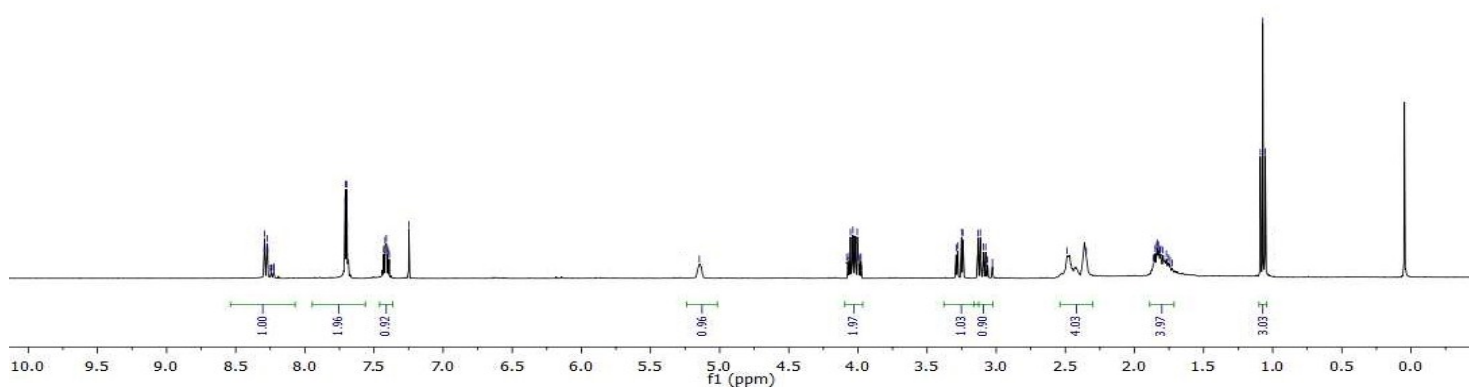
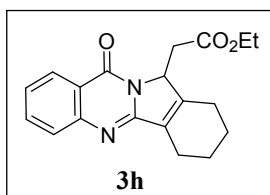


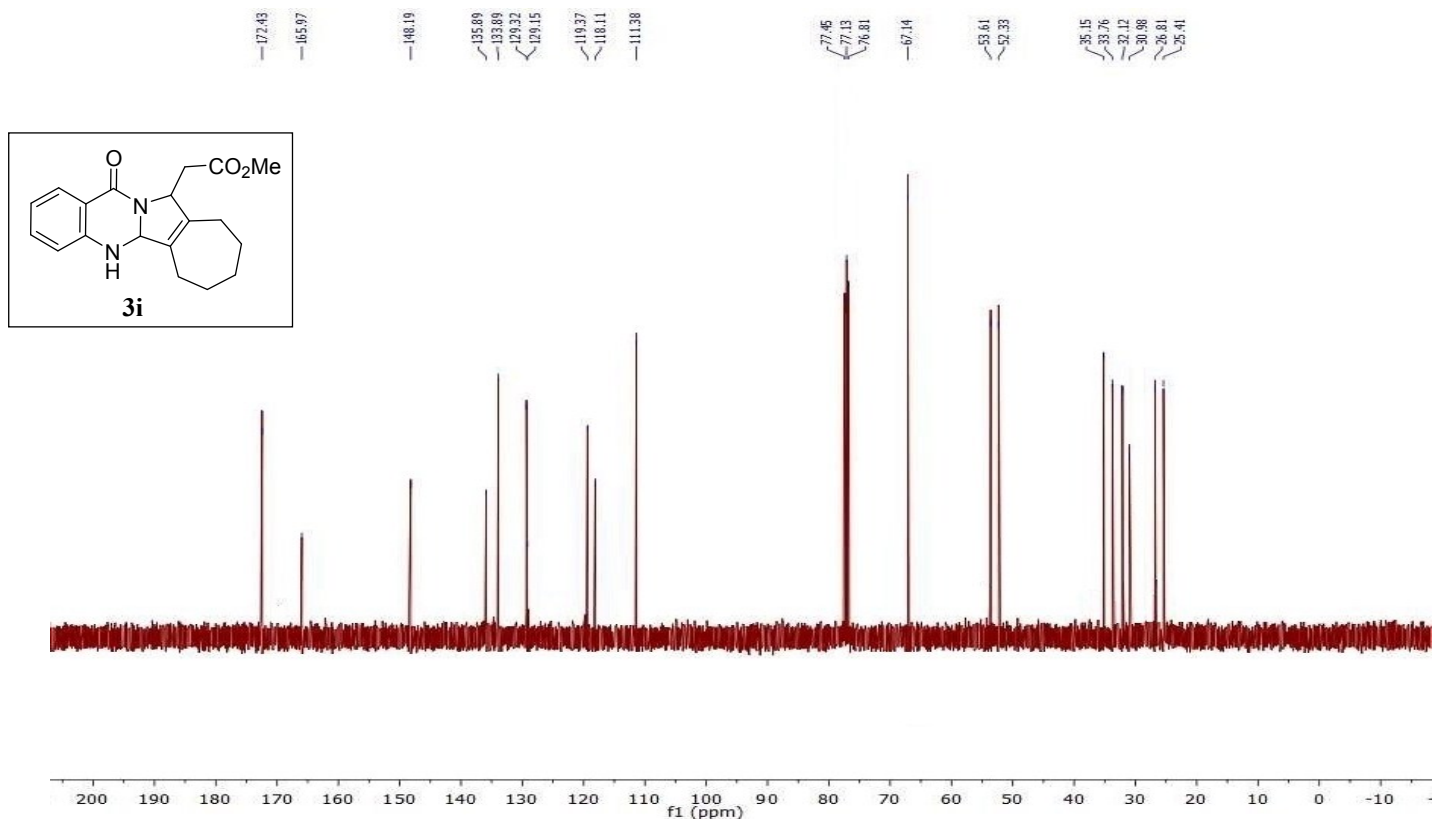
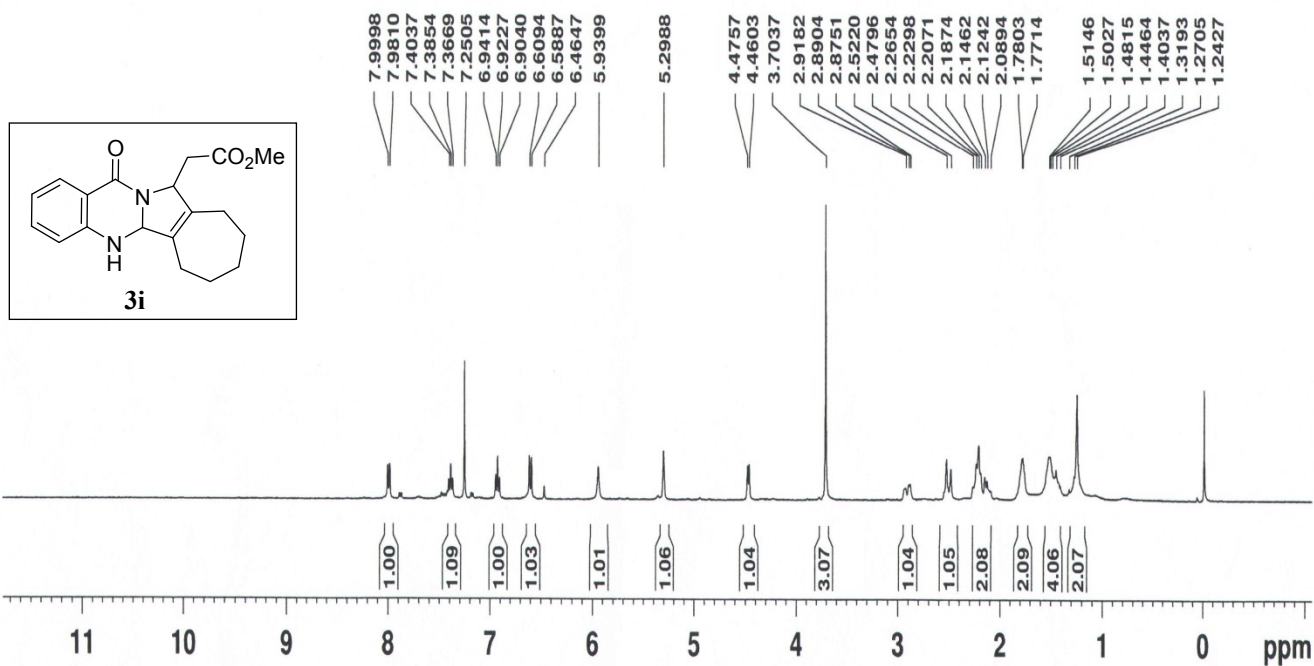


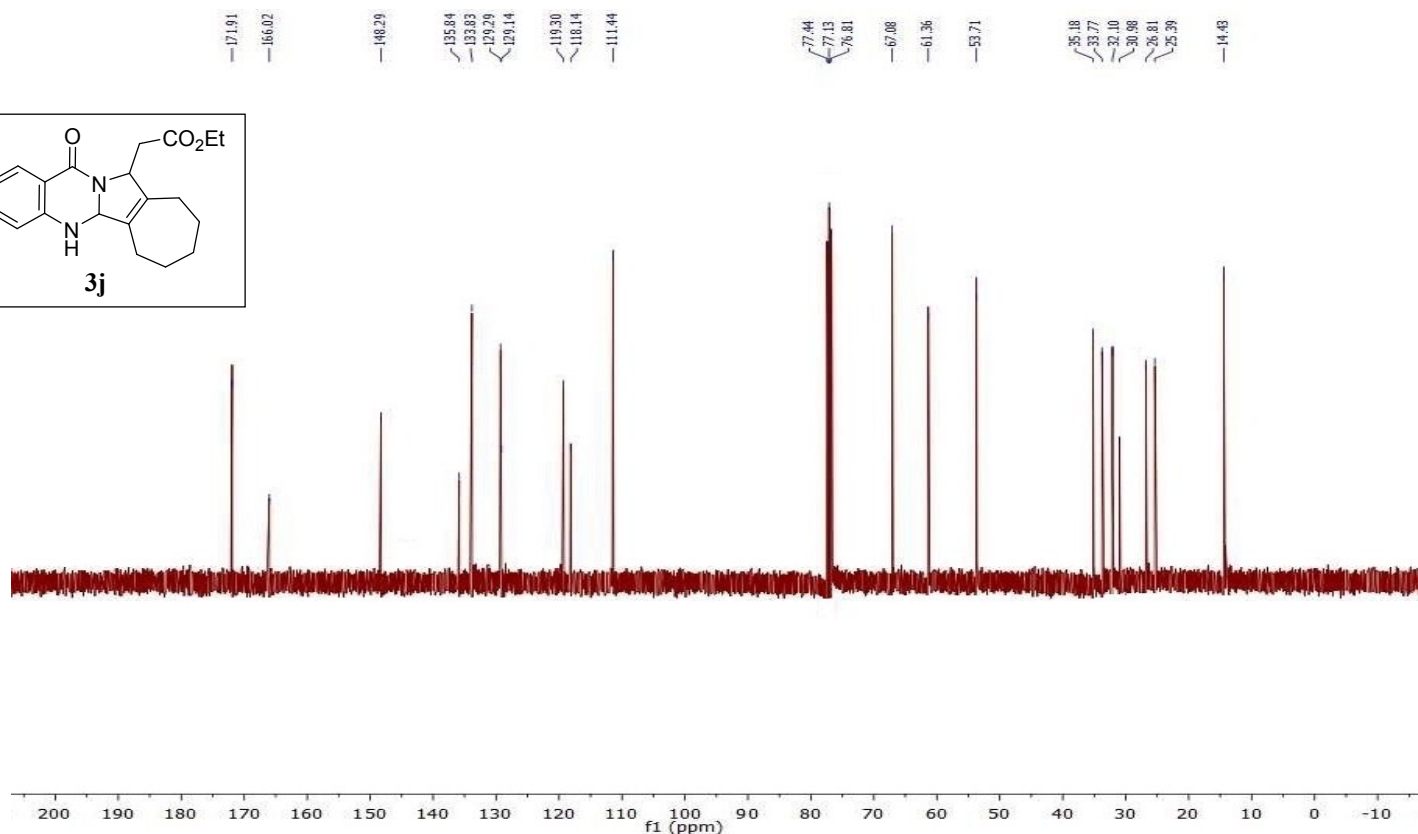
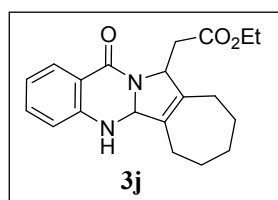
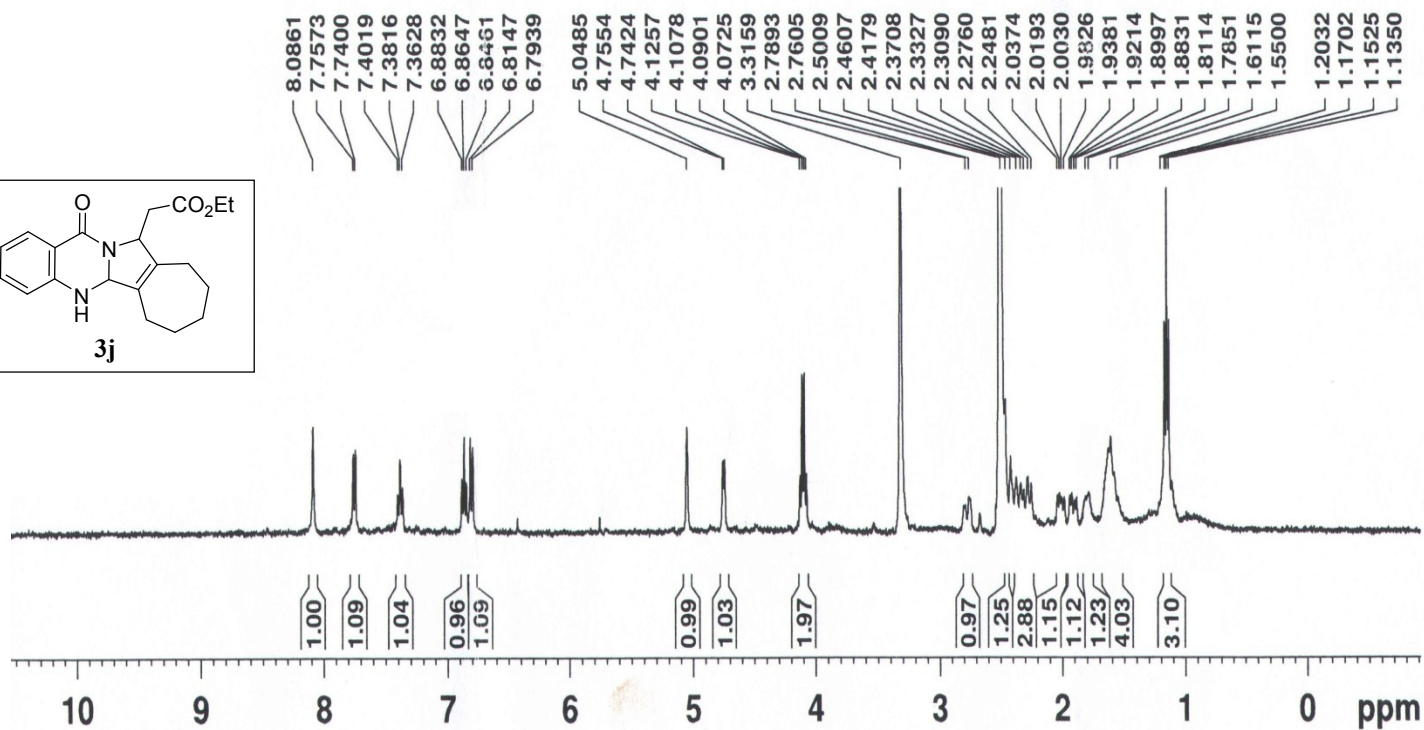
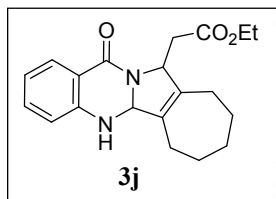


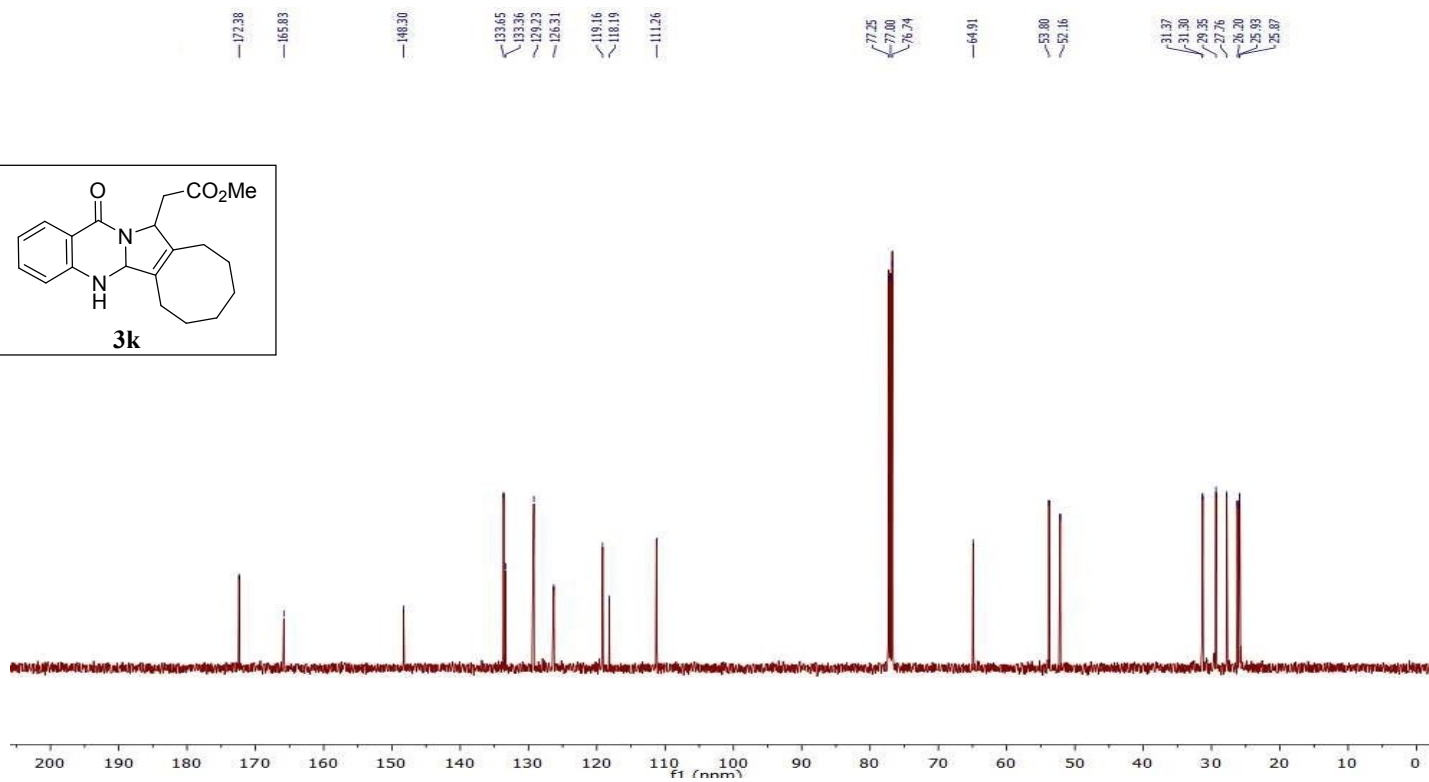
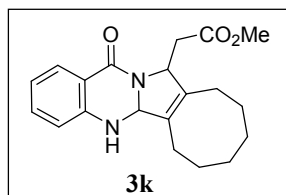
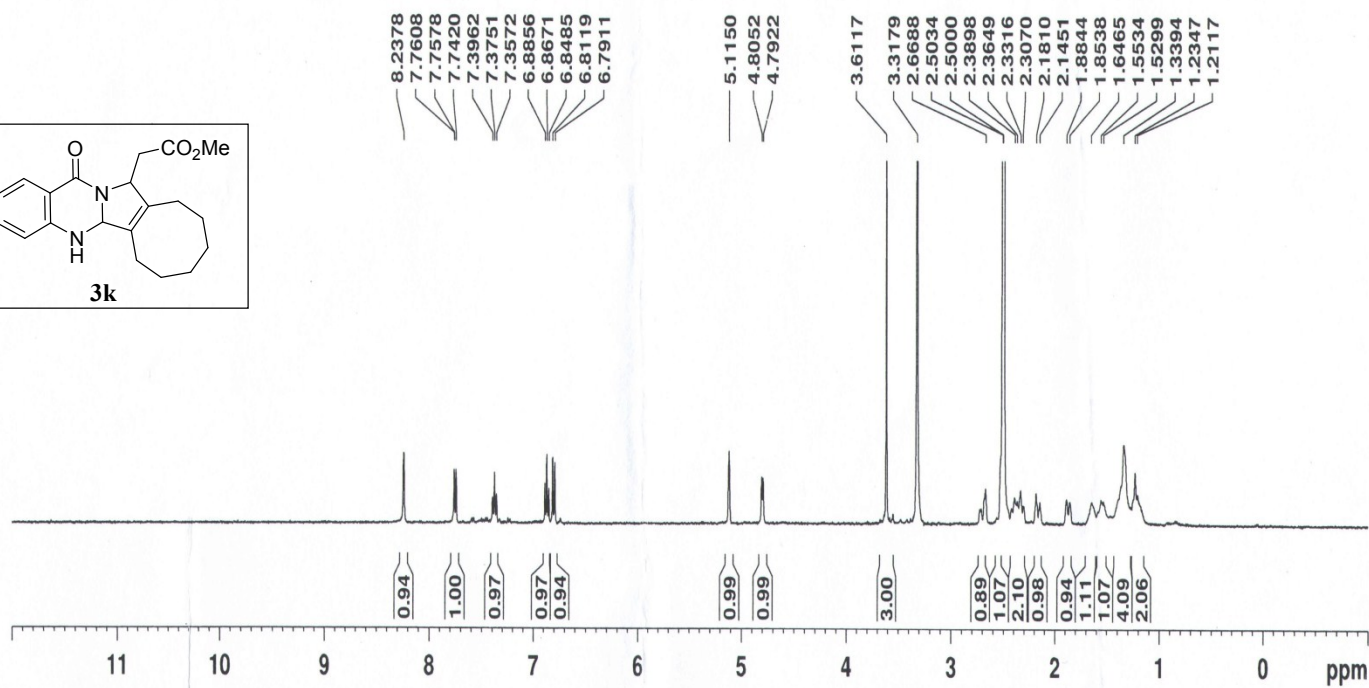
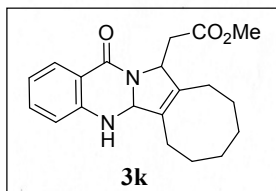


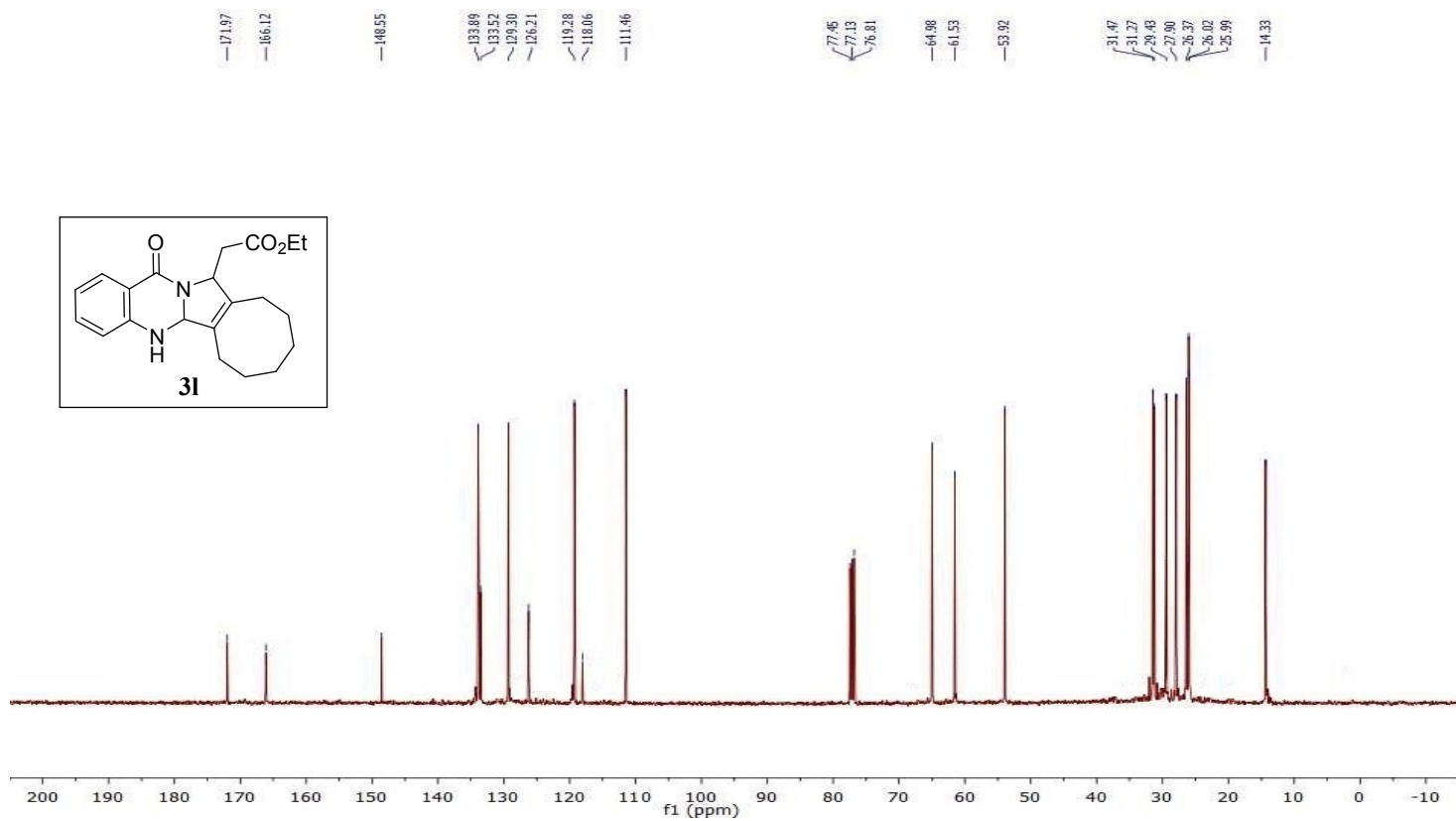
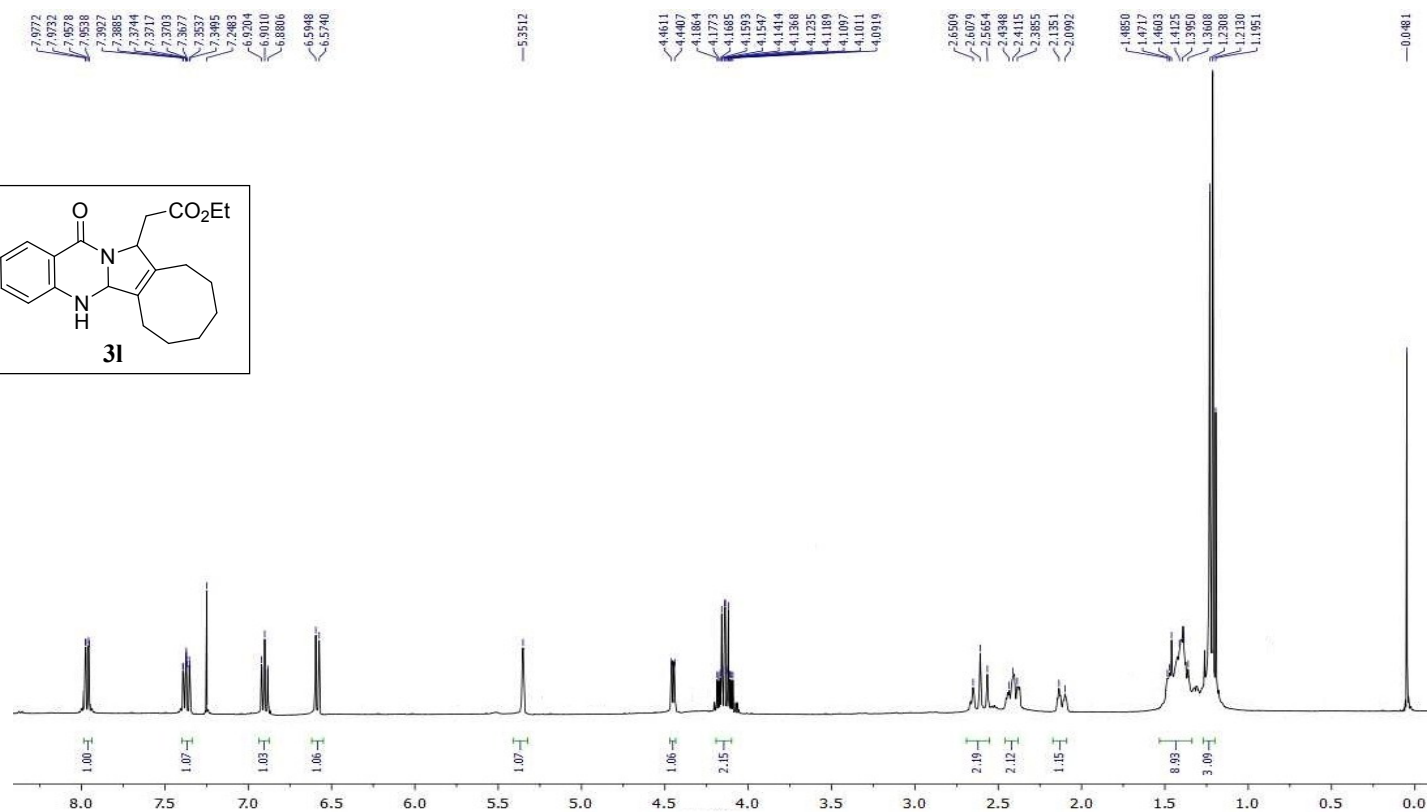
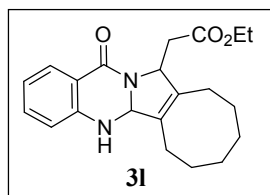


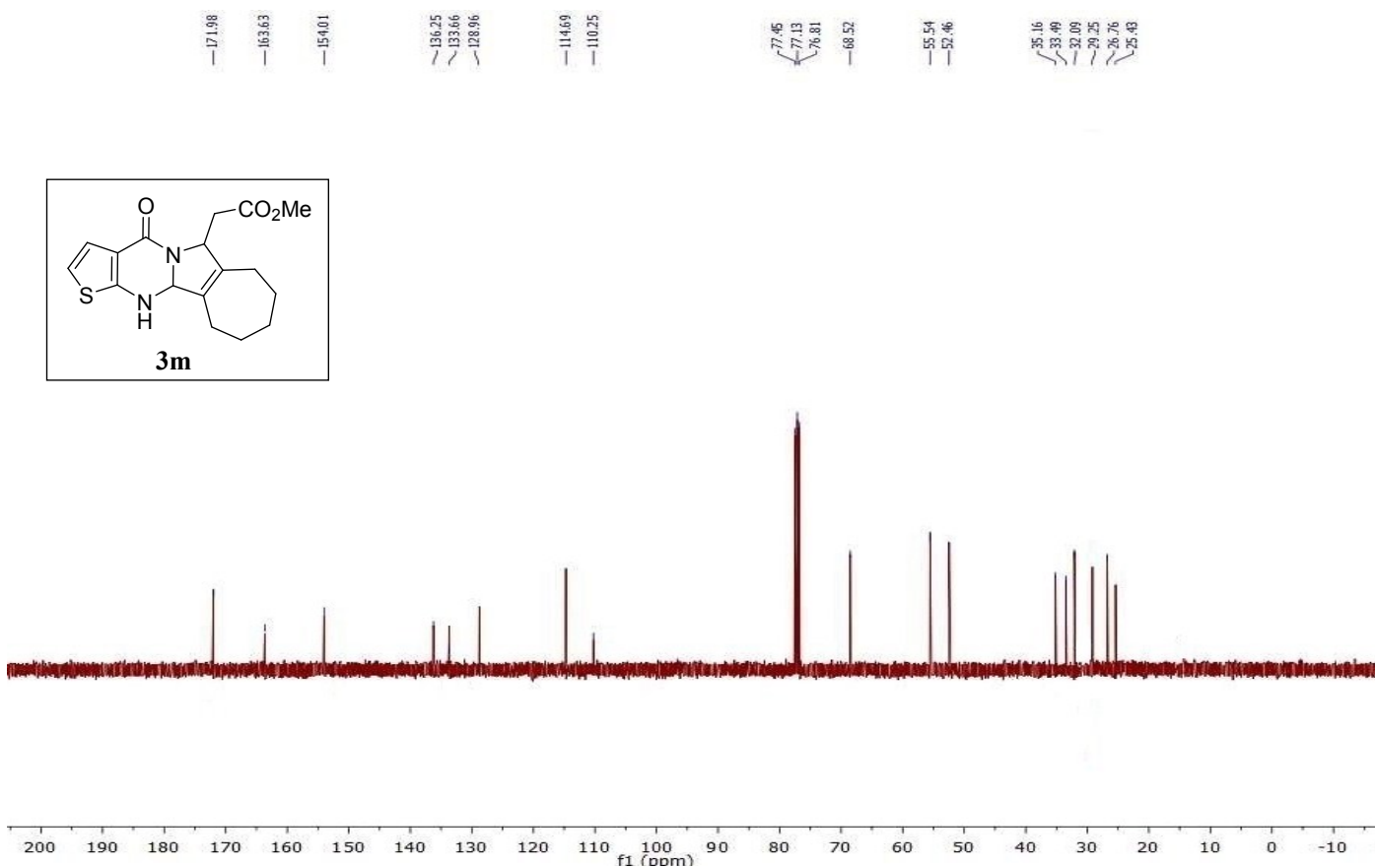
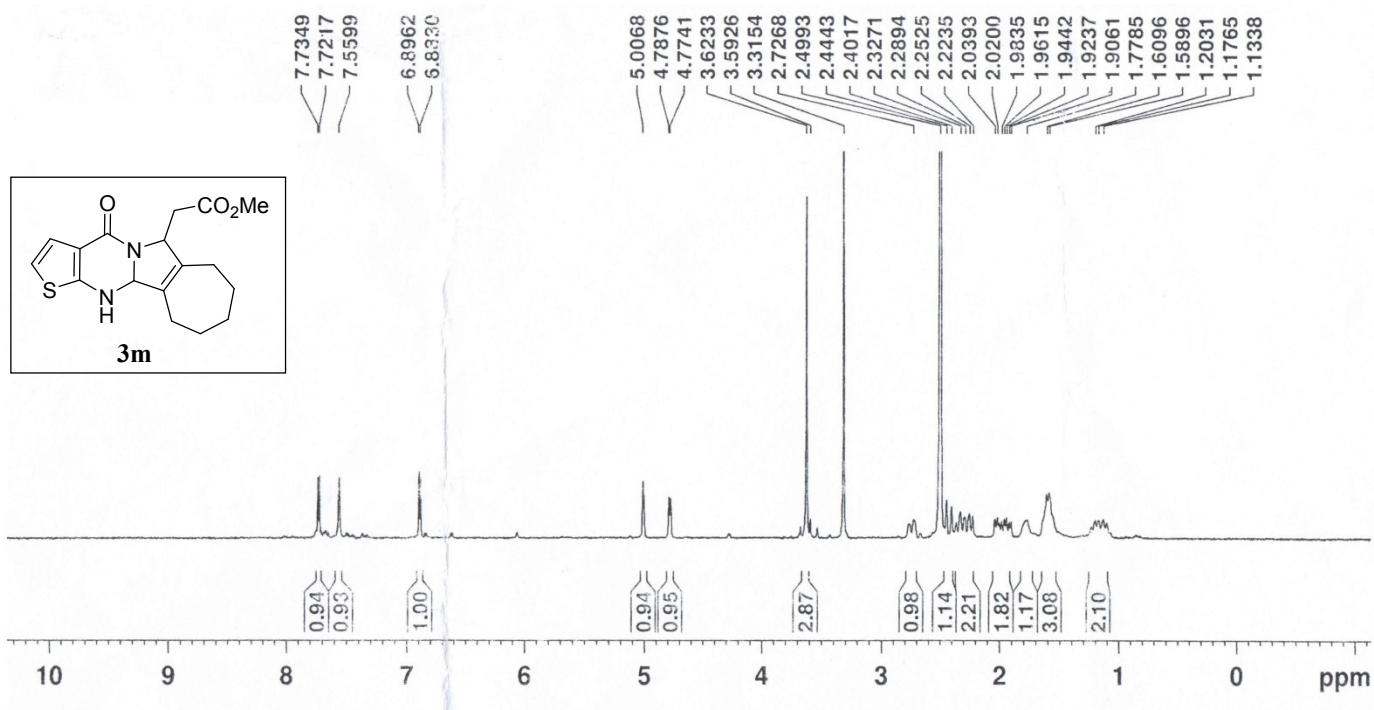


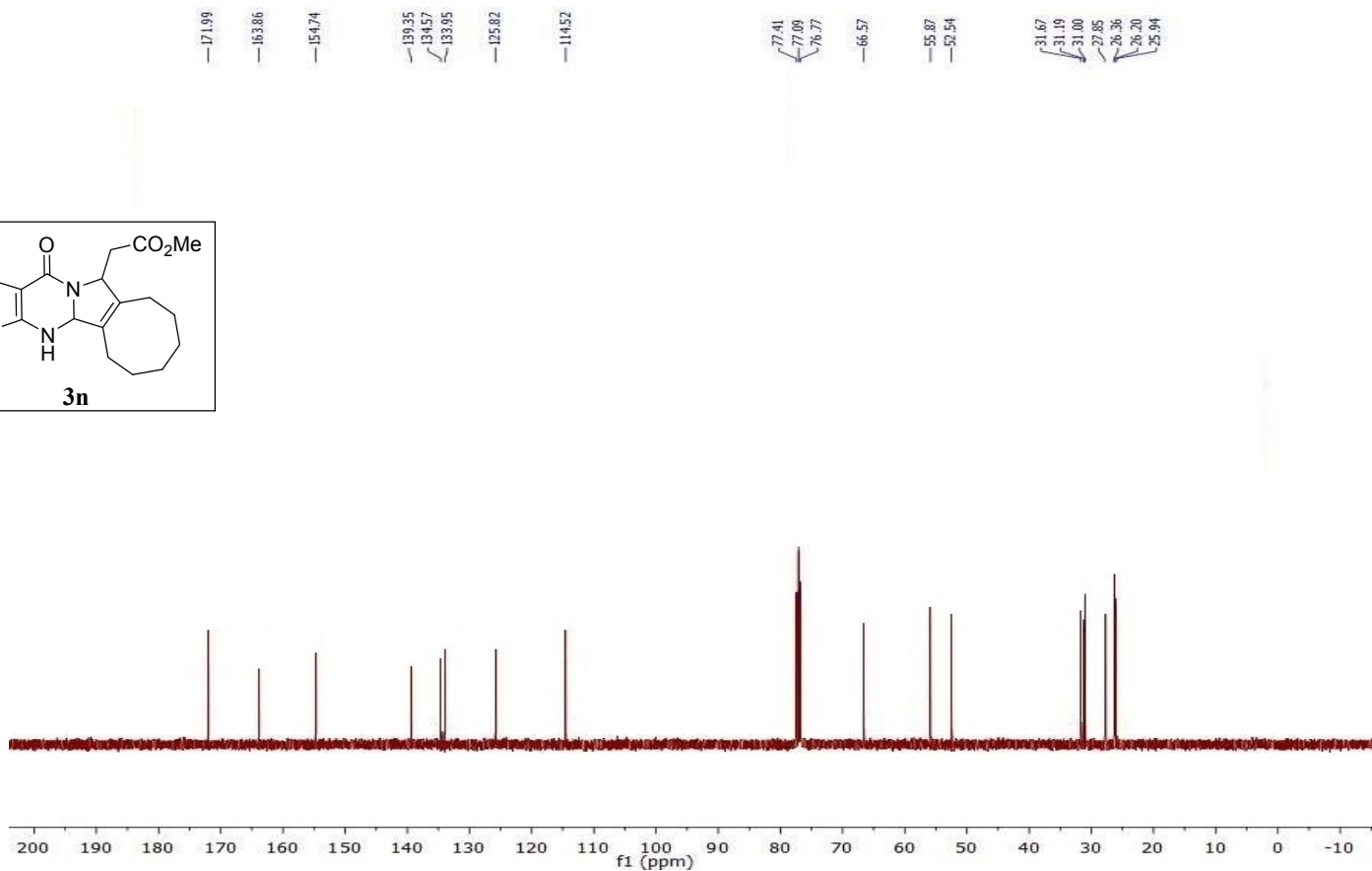
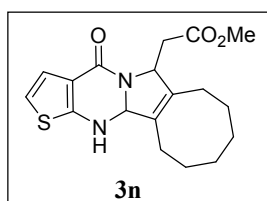
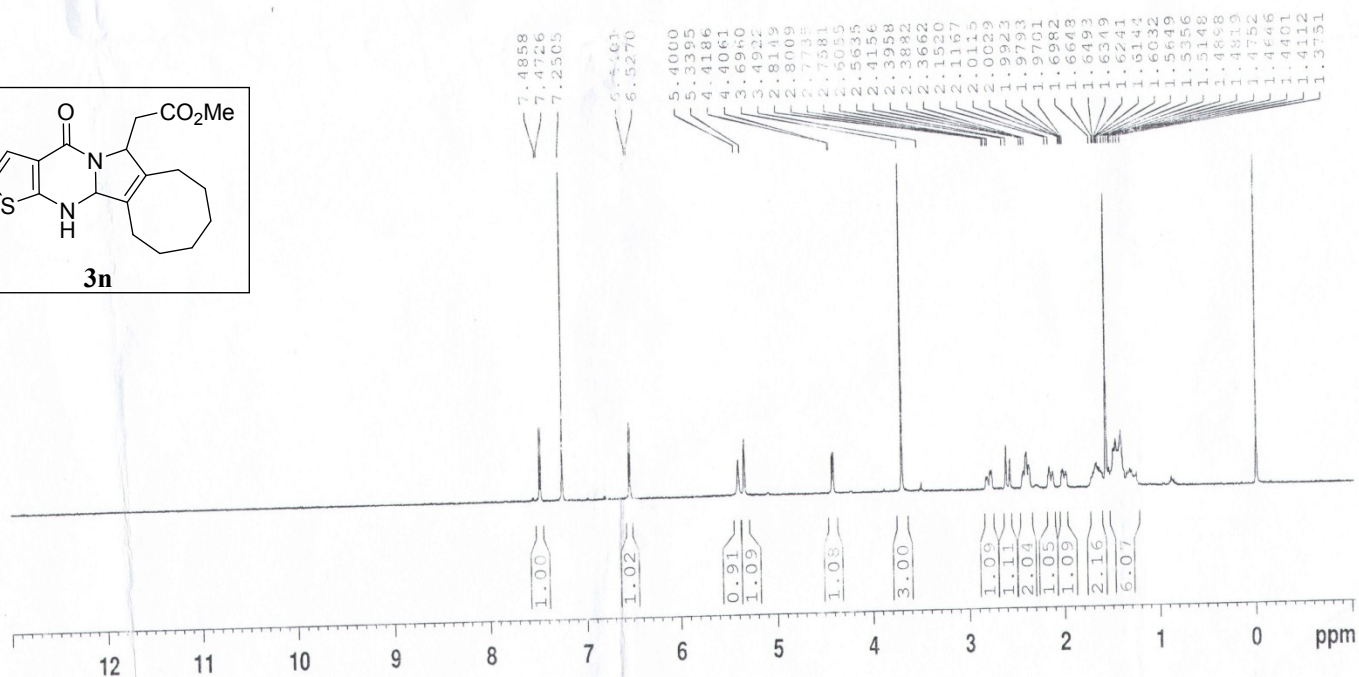
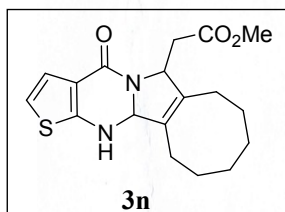


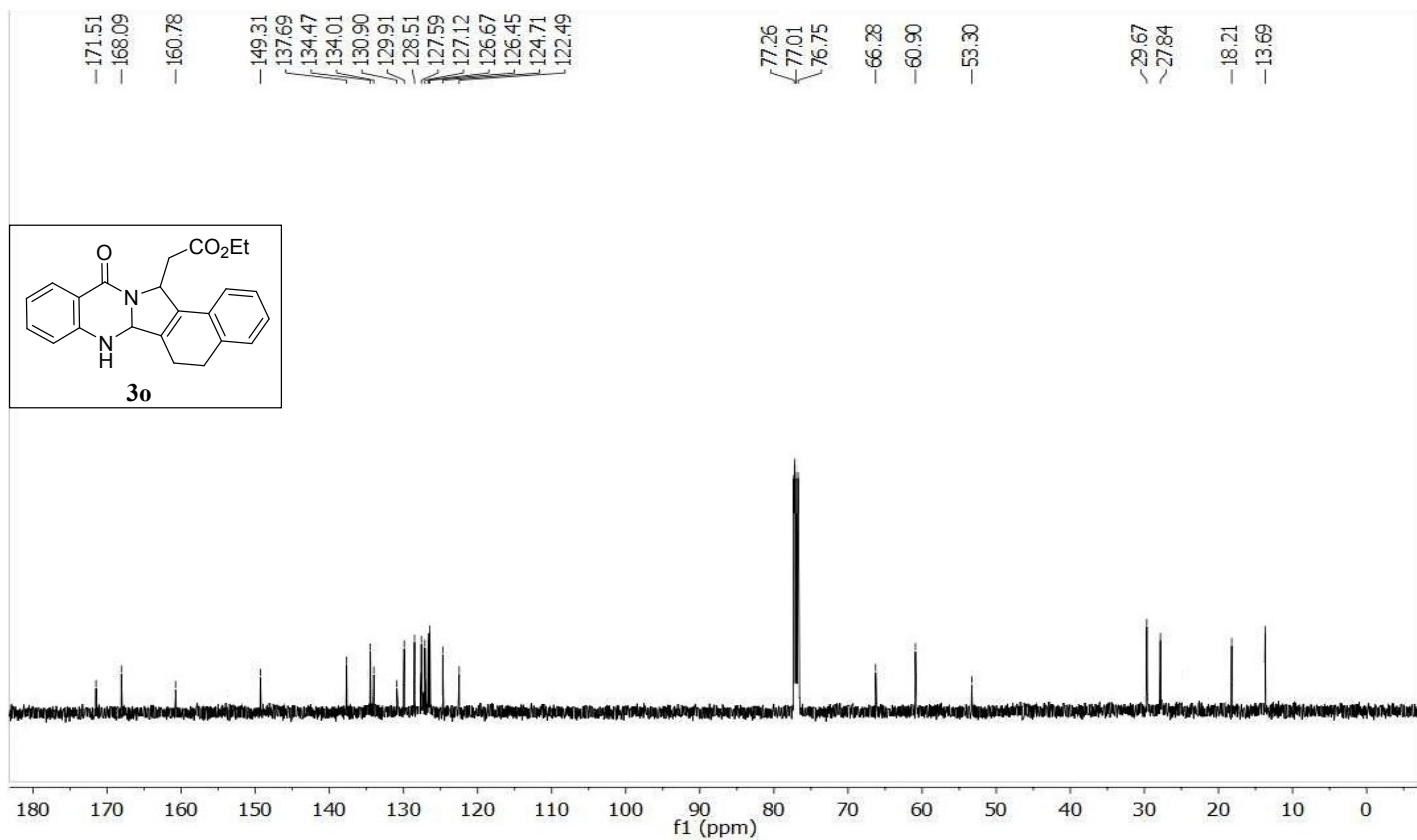
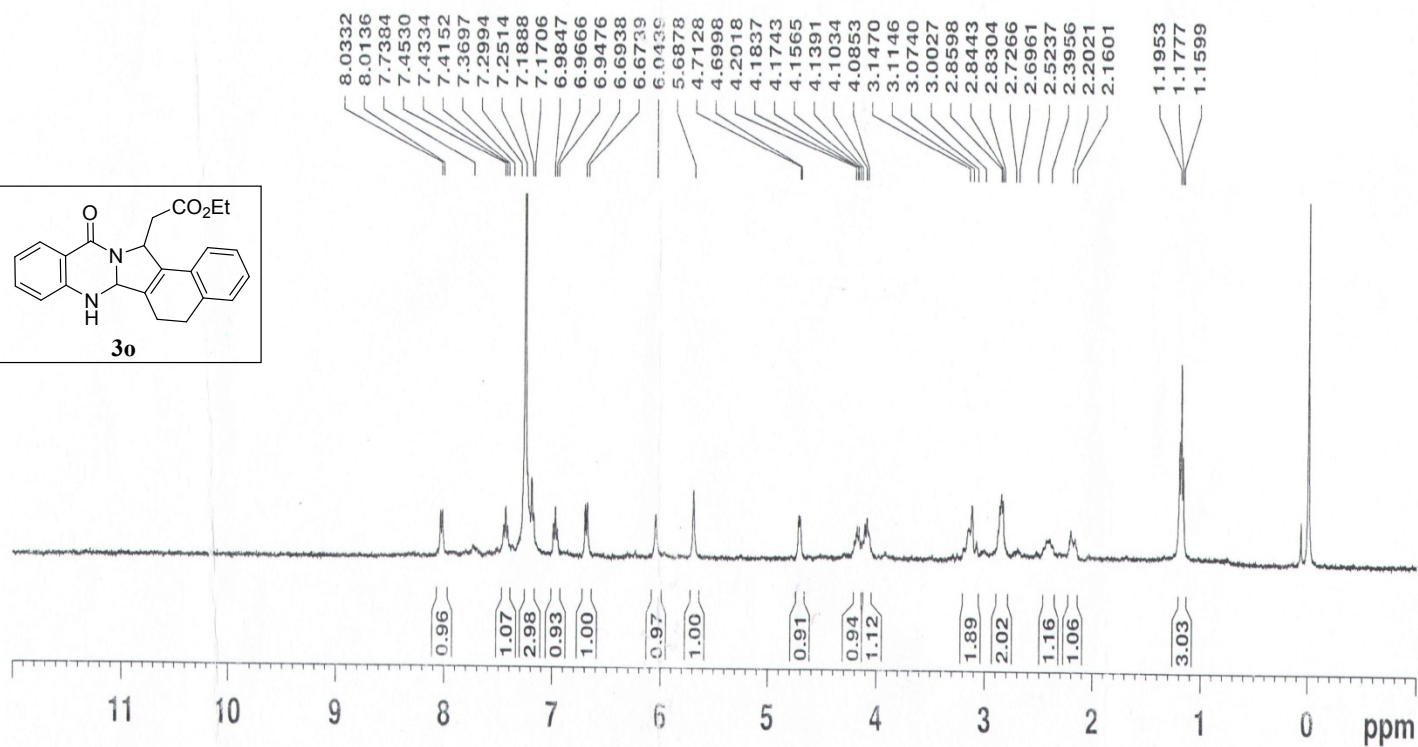
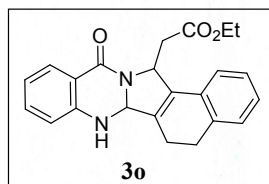


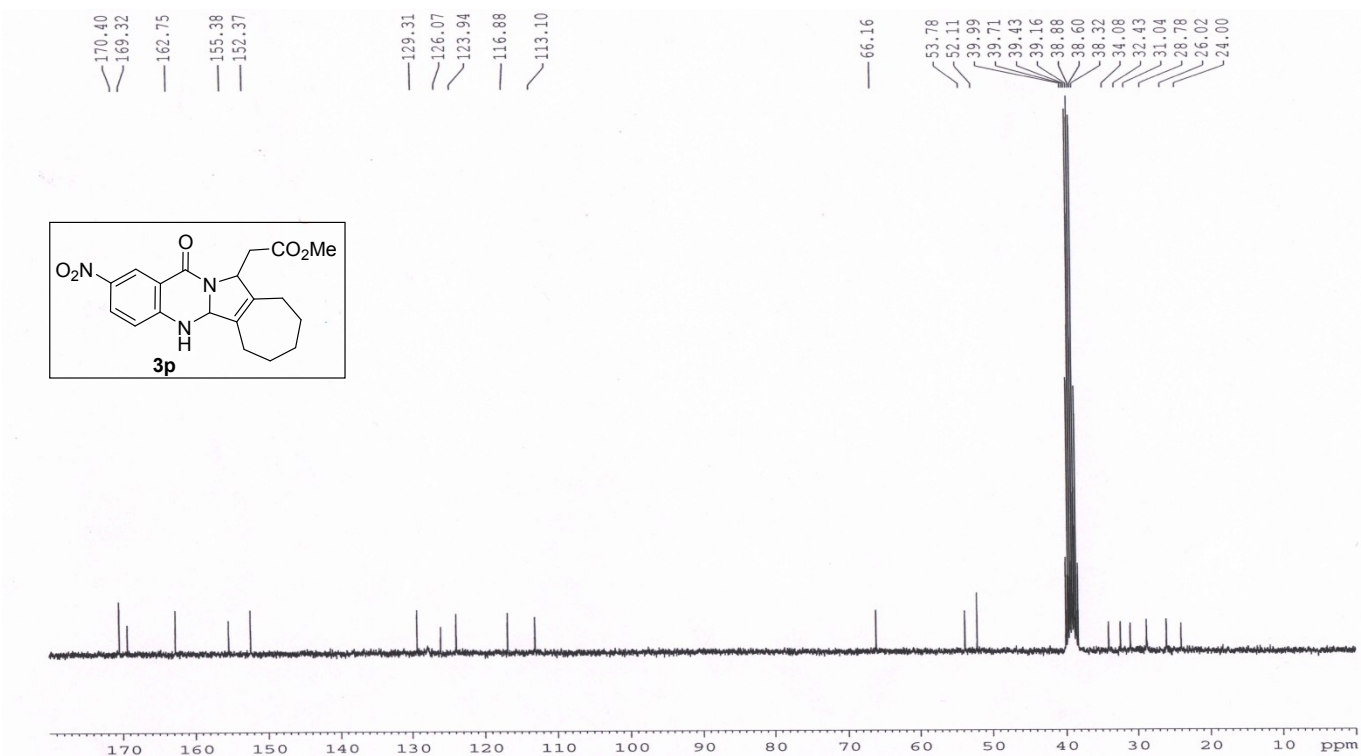
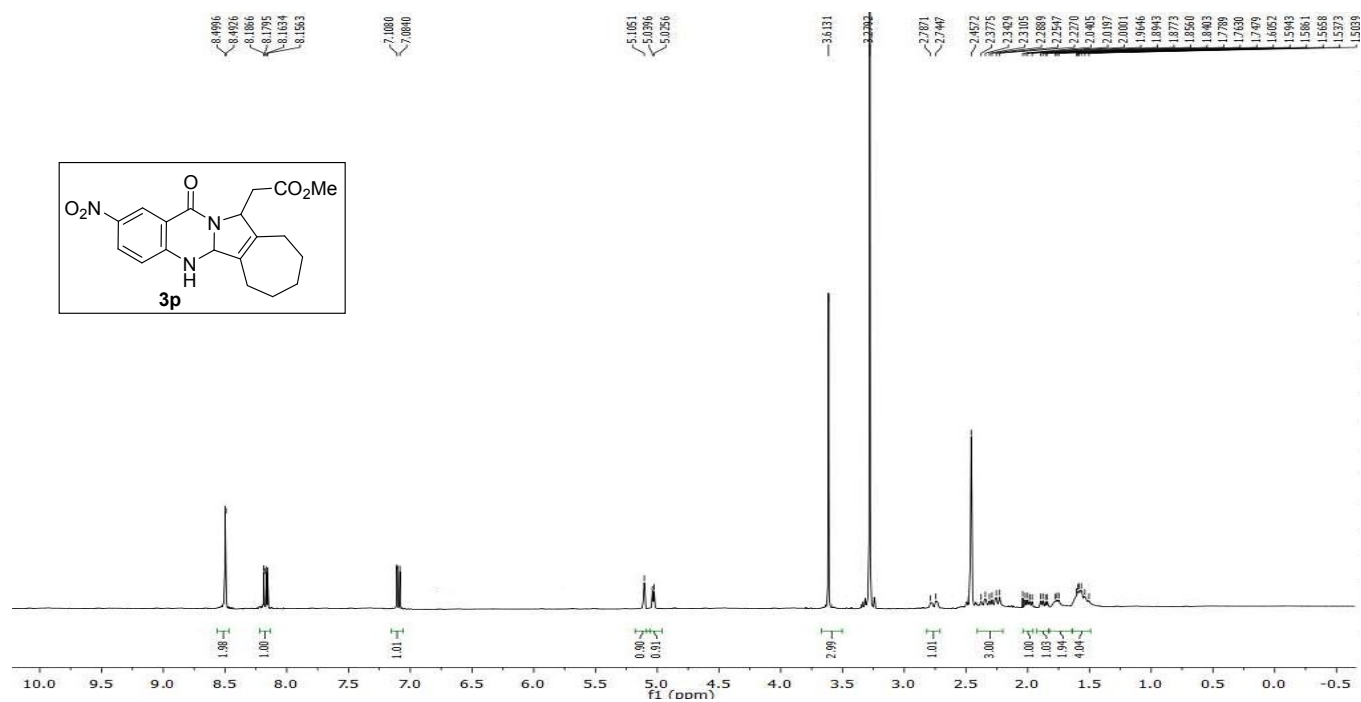


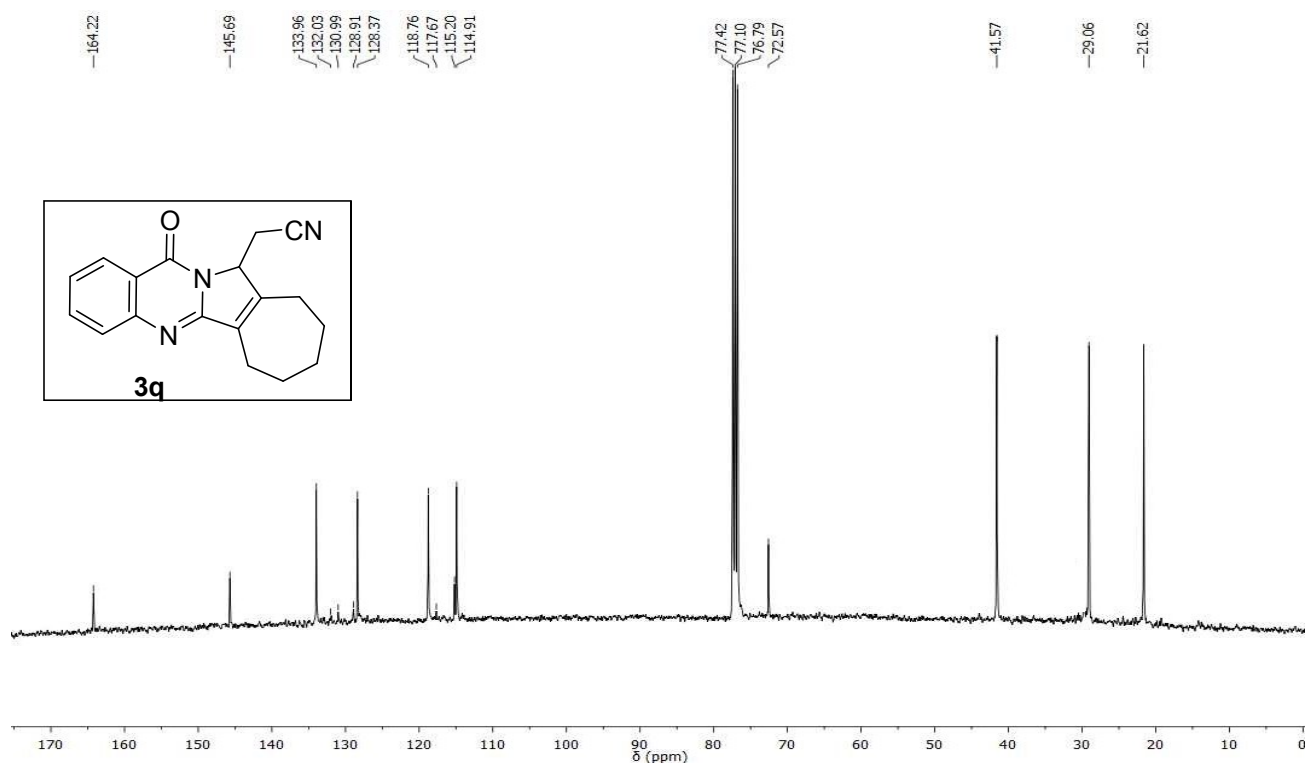
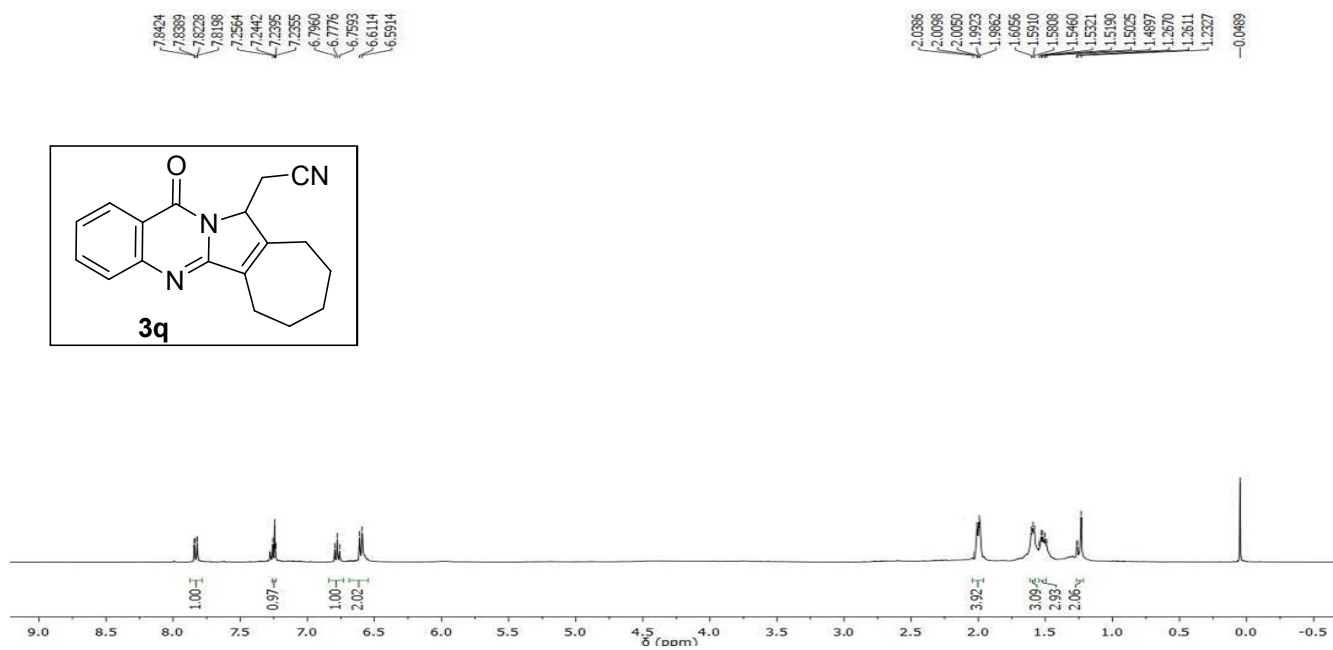
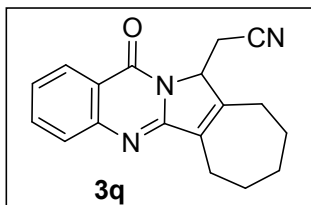


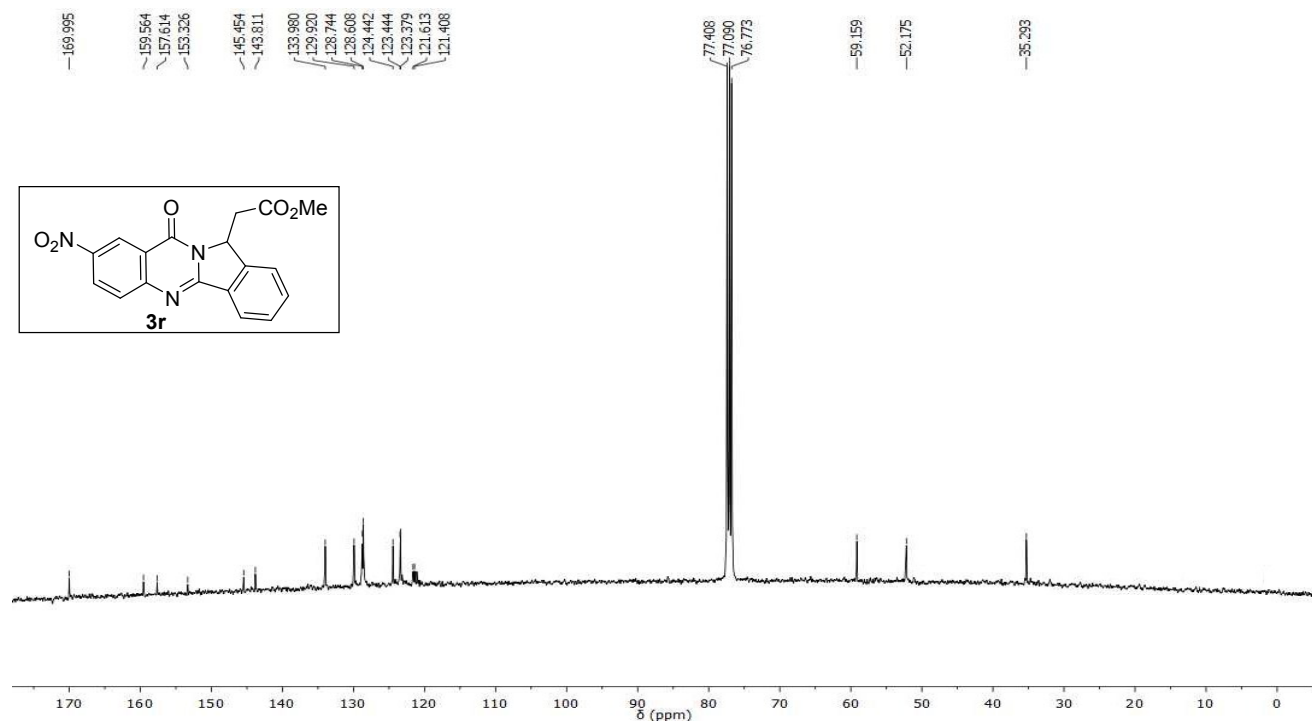
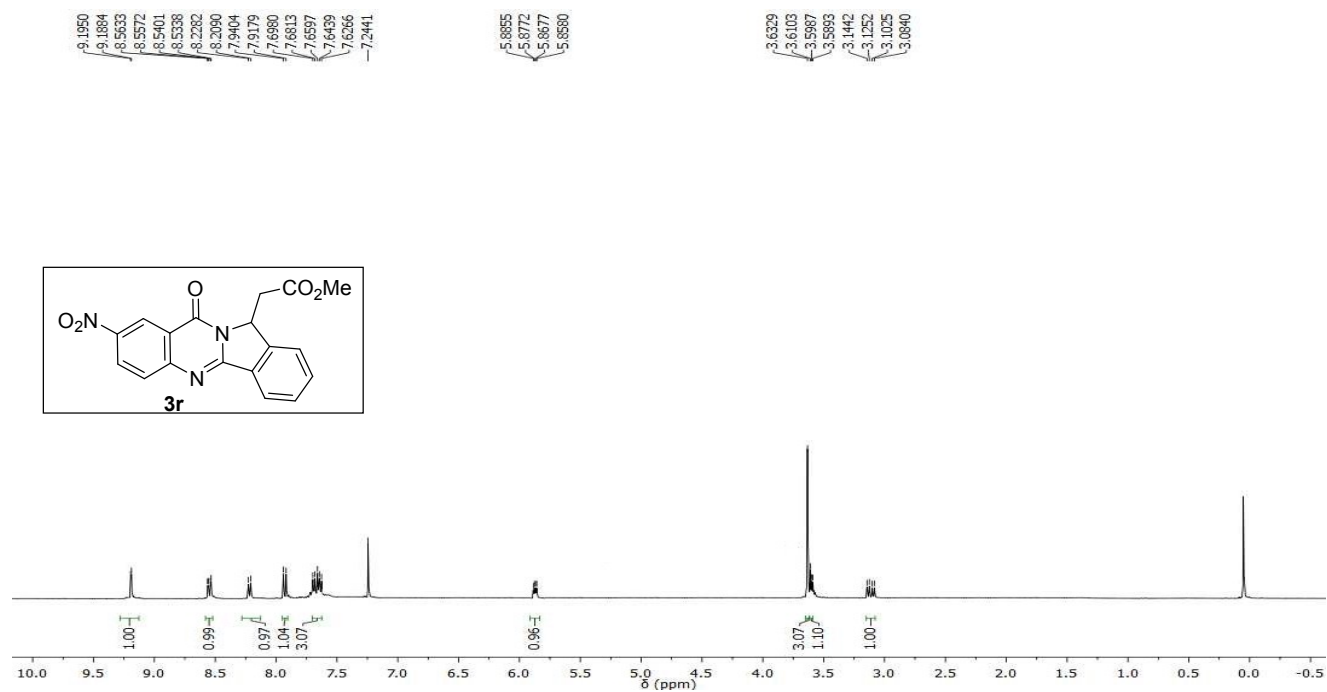


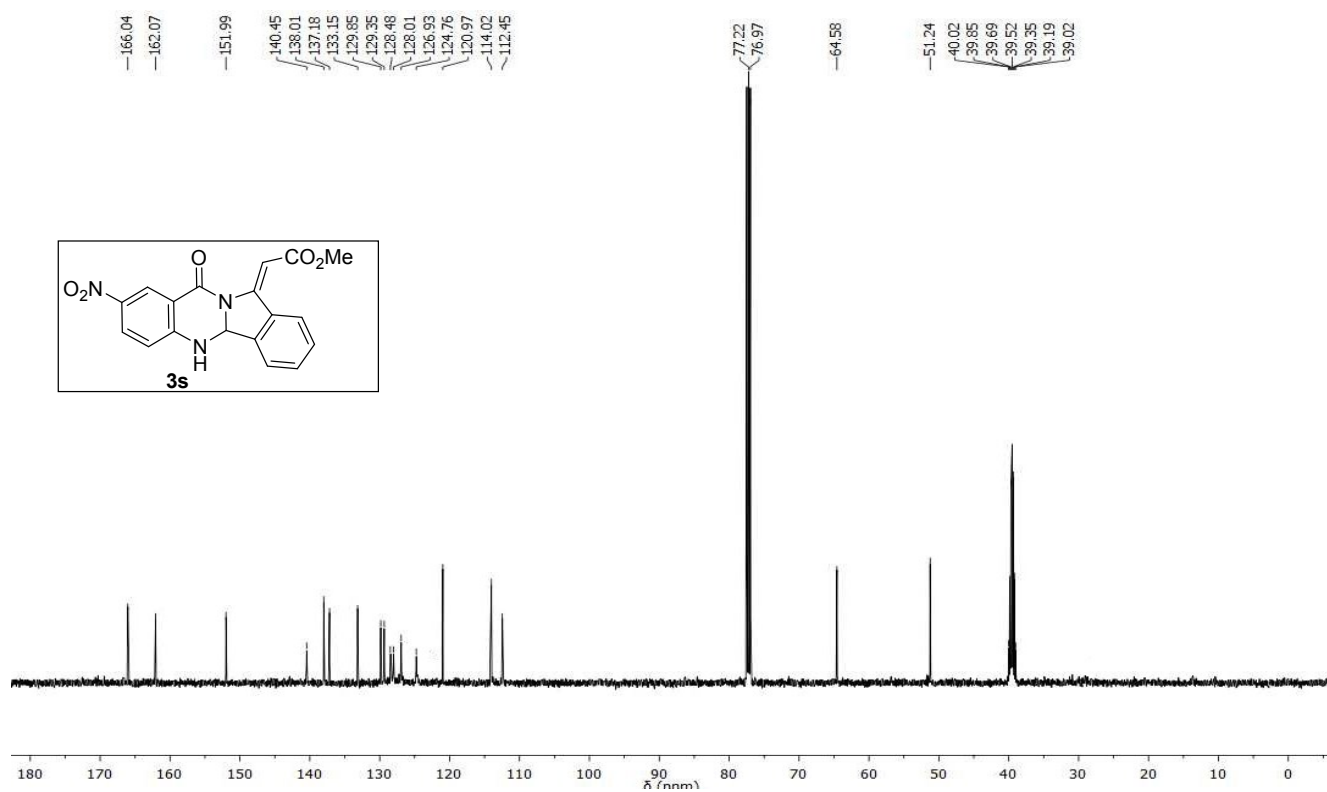
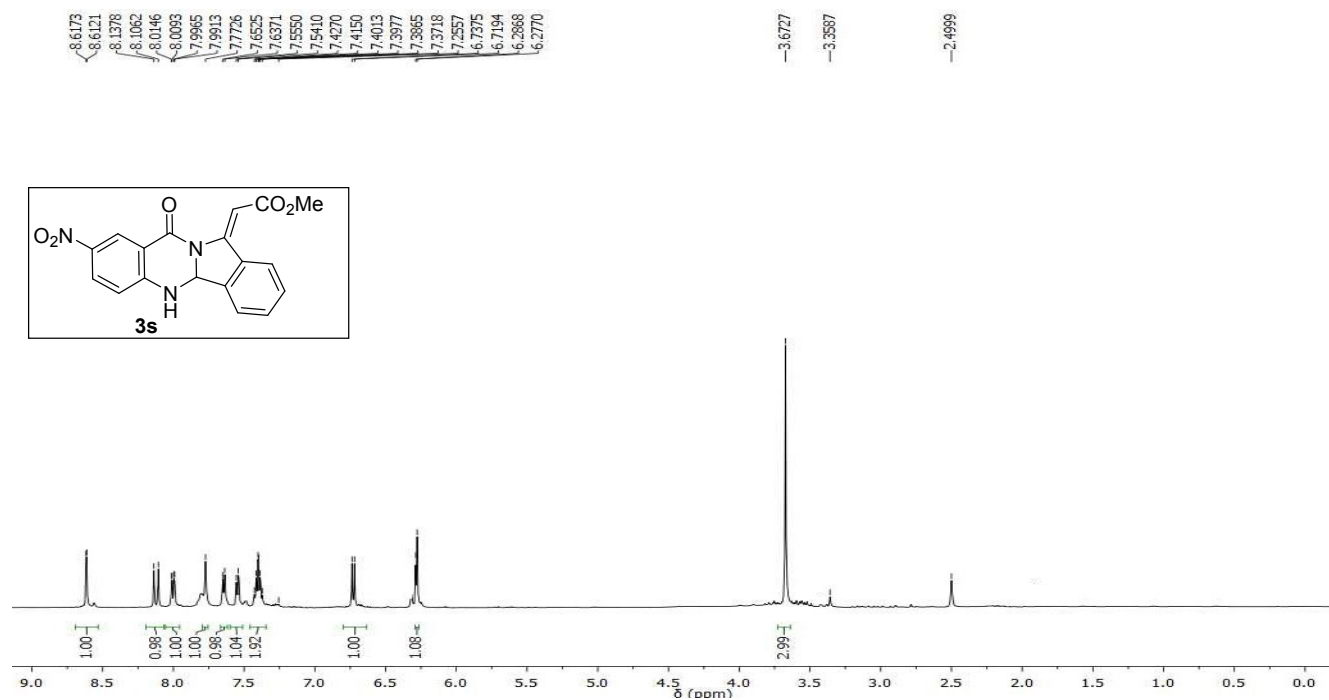


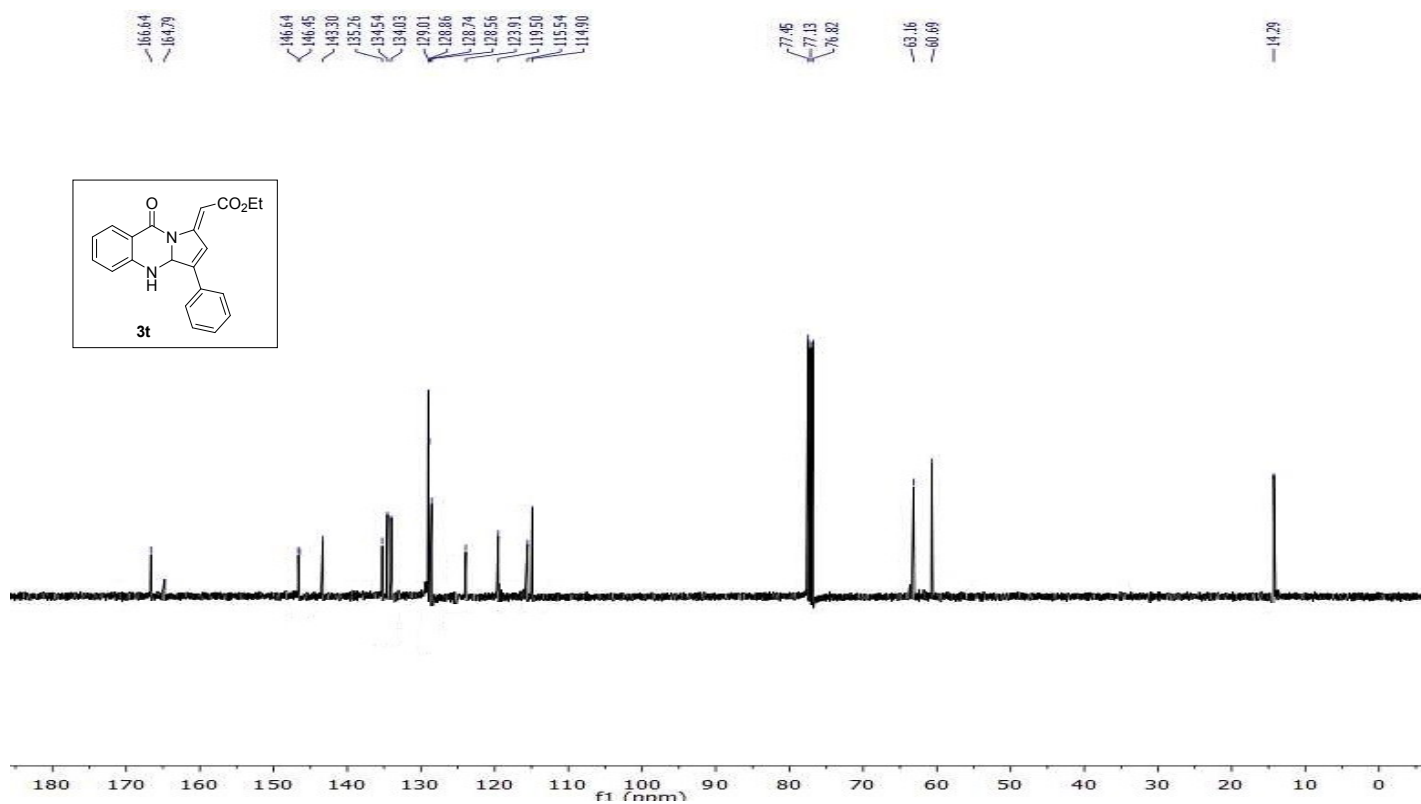
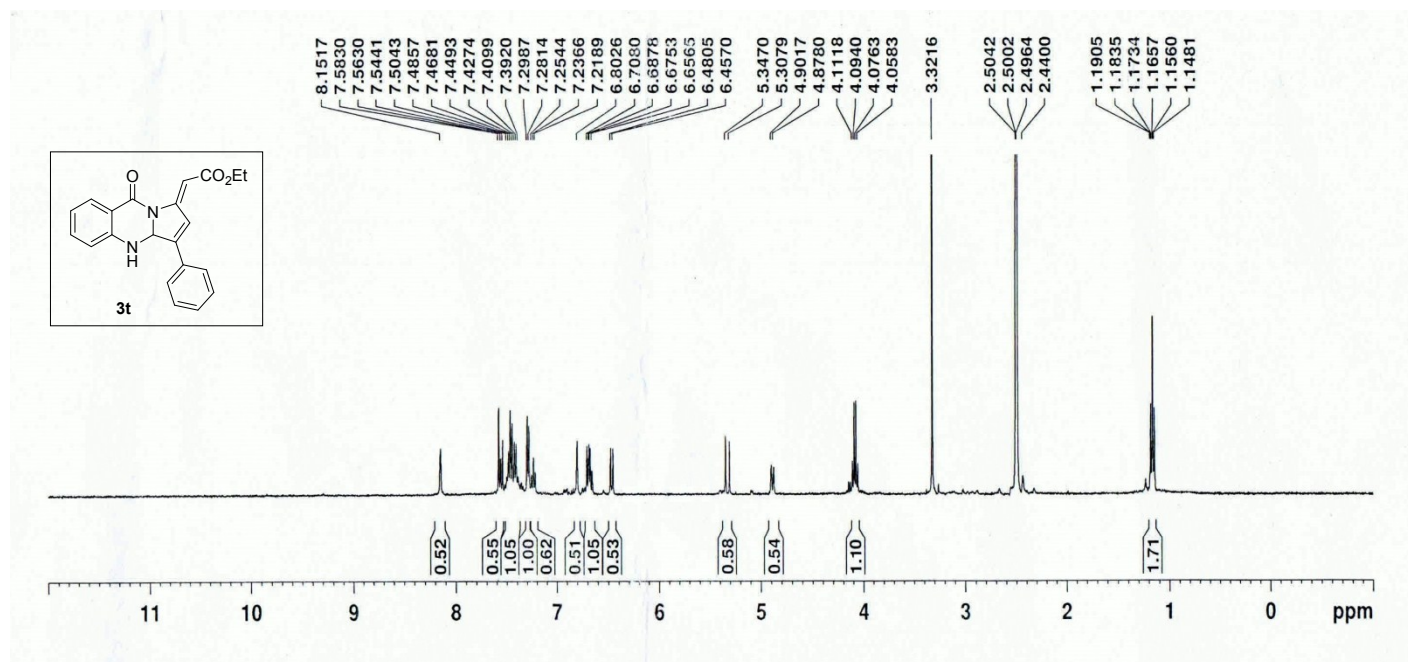












References:

1. (a) S. K. Manna, A. Mandal, S. K. Mondal, A. K. Adak, A. Jana, S. Das, S. Chattopadhyay, S. Roy, S. K. Ghorai, S. Samanta, M. Hossain and M. Baidya, *Org. Biomol. Chem.*, 2015, **13**, 8037; (b) S. K. Manna, S. K. Mondal, A. Ahmed, A. Mandal, A. Jana, M. Ikbal, S. Samanta and J. K. Ray, *RSC Adv.*, 2014, **4**, 2474; (c) S. K. Mondal, S. K. Manna, A. Mandal, S. Samanta and J. K. Ray, *Tetrahedron Lett.*, 2014, **55**, 6411; (d) A. Jana, S. K. Manna, S. K. Mondal, A. Mandal, A. Jana, B. K. Senapati, M. Jana and S. Samanta, *Tetrahedron Lett.*, 2016, **57**, 3722; (e) S. K. Mondal, A. Mandal, S. K. Manna, Sk. A. Ali, M. Hossain, V. Venugopal, A. Jana and S. Samanta, *Org. Biomol. Chem.*, 2017, **15**, 2411; (f) S. K. Mondal, S. A. Ali, S. K. Manna, A. Mandal, B. K. Senapati, M. Hossain and S. Samanta, *ChemSelect.*, 2017, **2**, 9312.
2. Y. Zheng, W-B. Song, S-W. Zhang and L-J. Xuan, *Org. Biomol. Chem.*, 2015, **13**, 6474.