

Supporting Information

## **Asymmetric catalytic concise synthesis of 3-(3-indolomethyl)oxindoles for the construction of trigolutes analogs**

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# Content

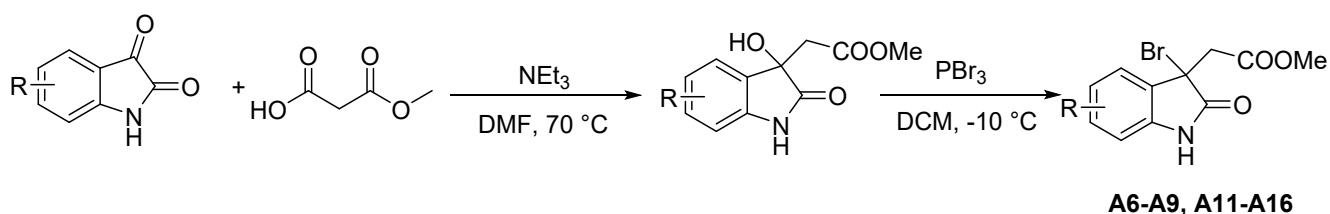
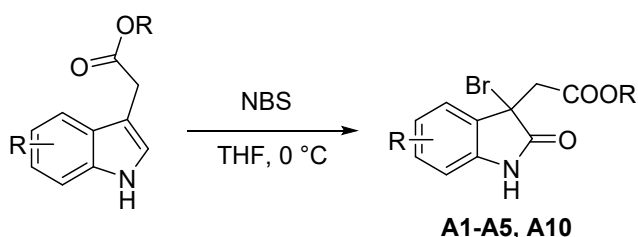
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## 1 General information

NMR characterization data were collected on Bruker ASCEND™ 400M and 600M.  $^1\text{H}$  NMR and  $^{13}\text{C}\{^1\text{H}\}$  NMR: chemical shifts  $\delta$  were recorded in ppm relative to tetramethylsilane and internally referenced to the residual solvent signal. Data were reported as follows: chemical shift, multiplicity (s = singlet, d = doublet, t = triplet, q = quartet, dd = doublet of doublets, td = triplet of doublets, dt = doublet of triplets, ddd = doublet of doublet of doublets, m = multiplet), coupling constants (Hz), integration. Ultra Performance Convergence Chromatography (UPC<sup>2</sup>) was performed on using Daicel Chiralcel IA-3, IC-3, ID-3, IE-3, IG-3, IH-3, OD-3, OX-3, OZ-3, AD-3. at 35 °C with UV detector at 211 nm, and enantiomeric excesses were determined in comparison with the authentic racemates. High resolution mass spectra (HRMS) were performed on Thermo Q-Exactive Focus (FTMS+c ESI) and data were reported as (m/z). Infrared spectra (IR) were recorded on Bruker Tensor II spectrometer with Plantium ATR accessory and the peaks are reported as absorption maxima ( $\nu$ ,  $\text{cm}^{-1}$ ). Optical rotations were measured on Rudolph Research Analytic Automatic Polarimeter, and reported as follows:  $[\alpha]_{\text{D}}^{25}$  (c: g/100 mL, in  $\text{CH}_2\text{Cl}_2$ ,  $\lambda = 589 \text{ nm}$ ). Melting point ranges were determined on OptiMelt. X-ray crystallographic data were collected by a Bruker D8 Venture Photon II. The experiments requiring substrates 3-Bromo-3-substituted oxindoles<sup>[1]</sup> and chiral *N,N'*-dioxide ligands<sup>[2]</sup> were synthesized according to known procedures and purified by recrystallization prior to use. All of the starting materials including the metal salts were purchased from TCI, Aladdin, Adamas, Acros, Aldrich and other companies, and used without further purification. The 3Å MS and inorganic base was purchased from Acros and oven-dried by the muffle furnace for 4 h prior to use. All the solvents were pre-dried over appropriate desiccants, and distilled prior to use. other commercial reagents were used without further purification. Reactions were monitored using thin-layer chromatography (TLC) on GF254 silica gel. Visualization of the developed plates was performed under UV light (254 nm) or using iodine, cobalt thiocyanate or  $\text{KMnO}_4$ . The products were purified by flash column chromatography with silicycle 300-400 mesh silica gel.

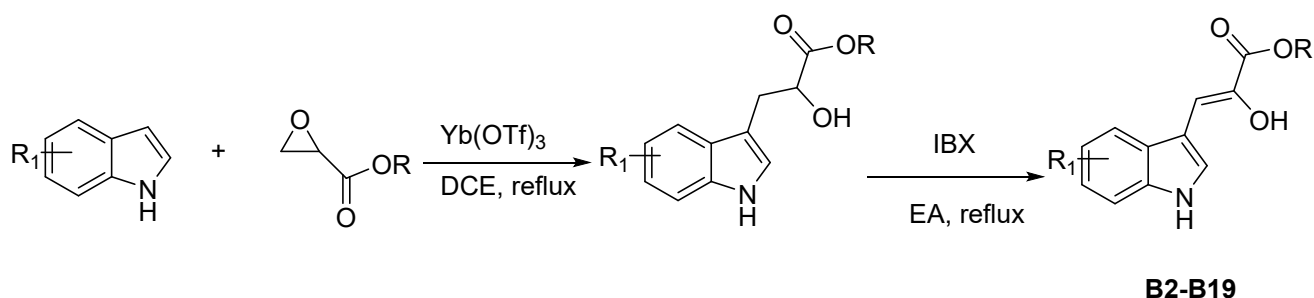
## 2 Typical procedure for preparation of substrates and products

### 2.1 Typical Synthesis Procedure of A and B



Synthesis Procedure of **A1-A5, A10**: *N*-bromosuccinimide (30.0 mmol, 5.34 g) was dissolved in cold THF (100 mL). The resulting THF solution was added dropwise via addition funnel to a stirring solution of 3-substituted indole (15 mmol) in *t*BuOH (100 mL) at room temperature over a period of 1 hour. The mixture was stirred for an additional hour. The solution was concentrated under reduced pressure. The crude material was subjected to column chromatography on silica gel (elute: petroleum ether/ethyl acetate = 10/1, v/v) to obtain the product.

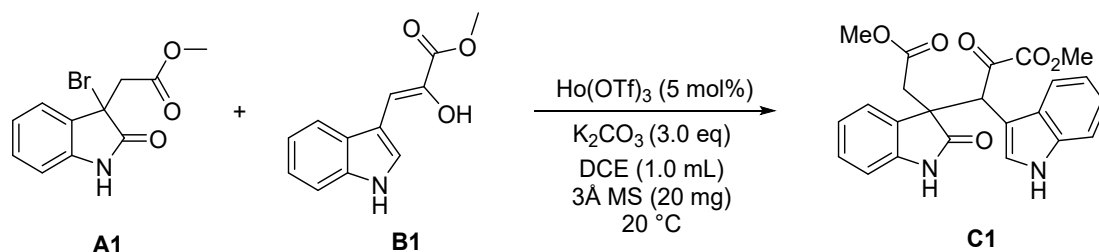
Synthesis Procedure of **A6-A9, A11-A16**: a solution of isatin (30 mmol) was dissolved in DMF and the corresponding acid (33 mmol) was added at room temperature, followed by addition of triethylamine (20 mol%). The mixture was stirred at 70 °C for 12 h and then washed with H<sub>2</sub>O. The aqueous layer was extracted with EA in three times, concentrated under reduced pressure, and then washed with DCM to obtain the crude product. The crude product (10 mmol) was dissolved in 10 mL DCM at -10 °C, followed by addition of PBr<sub>3</sub> (10 mmol), and monitored by TLC. The reaction mixture was quenched by with H<sub>2</sub>O, and extracted with DCM. The combined organic extracts were dried over Na<sub>2</sub>SO<sub>4</sub>, filtered and concentrated under reduce pressure. The residue was subjected to flash chromatograph on silica gel (petroleum ether/ethyl acetate = 10:1, v/v) and then recrystallized from the mixture of petroleum ether and ethyl acetate to afford 3-bromo-3-substituted oxindoles as a yellow solid.



**B1** was prepared according to previous literature.<sup>[3]</sup> Synthesis Procedure of **B2-B19**: Glycidic acid ester (4.0 g, 39.2 mmol) was added in one portion by syringe to a mixture of indole (78.4 mmol) and Yb(OTf)<sub>3</sub> (7.3 g, 11.7 mmol) in DCE (50 mL) under N<sub>2</sub>. The mixture was refluxed for 1.5 h. The reaction was quenched with sat. Na<sub>2</sub>CO<sub>3</sub> (50 mL) and the solution was acidified with 2M HCl (150 mL). The solution was extracted with CH<sub>2</sub>Cl<sub>2</sub> (3 x 50 mL) and the combined organic layers were washed with brine (50 mL), dried (Na<sub>2</sub>SO<sub>4</sub>), and concentrated under reduced pressure. The residue was subjected to flash chromatograph on silica gel (petroleum ether/ethyl acetate = 8:1, v/v) to give methyl -α-hydroxy-1H-indole-3-propanoate. Methyl -α-hydroxy-1H-indole-3-propanoate (10 mmol) was dissolved in

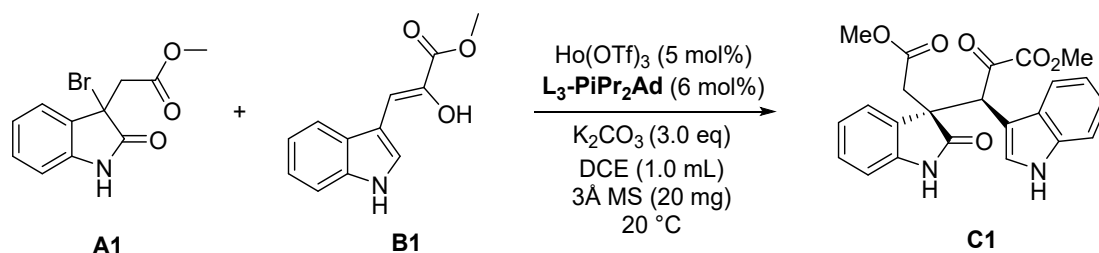
ethyl acetate (20 mL) in a round-bottomed flask equipped with a reflux condenser. Next, o-iodoxybenzoic acid (IBX) was added and the mixture was heated at reflux temperature (monitored by TLC). Upon completion, the reaction was cooled to room temperature and the crude mixture was passed through a pad of silica gel using ethyl acetate to rinse. Upon concentration, the residue was subjected to flash chromatograph on silica gel (petroleum ether/ethyl acetate = 8:1, v/v) and then recrystallized from the mixture of petroleum ether and ethyl acetate to afford the final product as a white solid.

### 2.3 Typical procedure for preparation of the racemic products



A dry reaction tube was charged with 3-Bromo-3-substituted oxindoles **A1** (42.6 mg, 0.15 mmol), methyl (Z)-2-hydroxy-3-(1H-indol-3-yl)acrylate **B1** (21.7 mg, 0.1 mmol),  $\text{K}_2\text{CO}_3$  (41.4 mg, 0.3 mmol, 3.0 equiv.), 3Å MS. (20.0 mg), and 1.0 mL DCE in the air. The reaction was performed at 20 °C for 19 hours. The reaction mixture was directly subjected to flash column chromatography on silica gel (eluent: petroleum ether/ethyl acetate = 2:1 to 1:1, v/v) to afford the corresponding products **C1**.

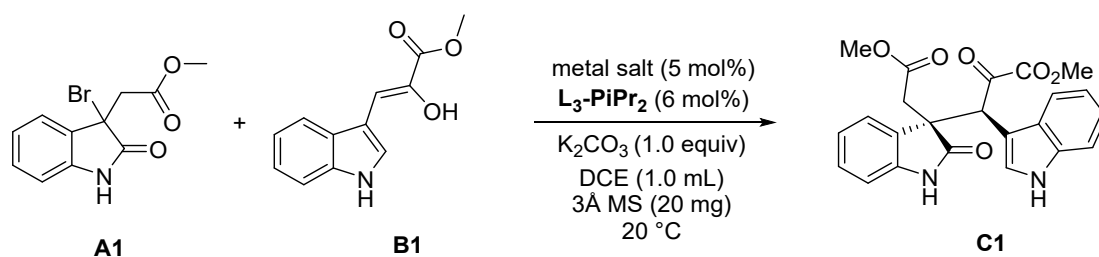
### 2.4 Typical procedure for preparation of the asymmetric products



An oven-dried test tube was charged with *N,N'*-dioxide **L3-PiPr2Ad** (5.5 mg, 0.006 mmol),  $\text{Ho(OTf)}_3$  (3.1 mg, 0.005 mmol), and 0.5 mL DCM under  $\text{N}_2$  atmosphere. After the mixture was stirred at 35 °C for 30 min, the solvent was removed in vacuo. 3-Bromo-3-substituted oxindoles **A1** (42.6 mg, 0.15 mmol), methyl (Z)-2-hydroxy-3-(1H-indol-3-yl)acrylate **B1** (21.7 mg, 0.1 mmol),  $\text{K}_2\text{CO}_3$  (41.4 mg, 0.3 mmol, 3.0 equiv.), 3Å MS. (20 mg), and 1.0 mL DCE were added in the glovebox. The reaction was performed at 20 °C for 11 hours. The reaction mixture was directly subjected to flash column chromatography on silica gel at -35°C (eluent: petroleum ether/ethyl acetate = 2:1 to 1:1, v/v) to afford the corresponding products **C1**. By the way, the product suffered somewhat epimerization after the chromatography on silica gel at room temperature.

### 3 Optimization of the reaction conditions

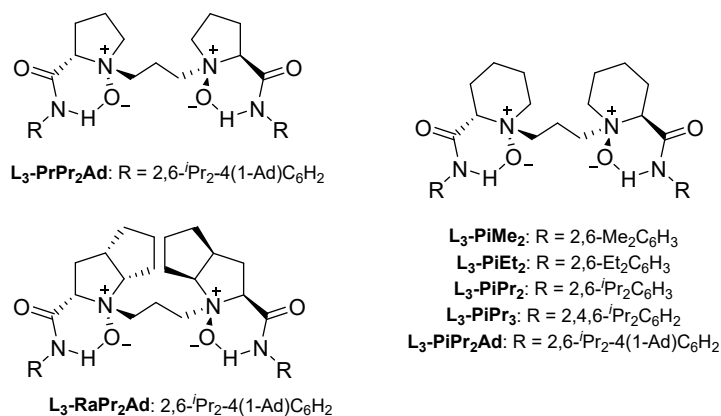
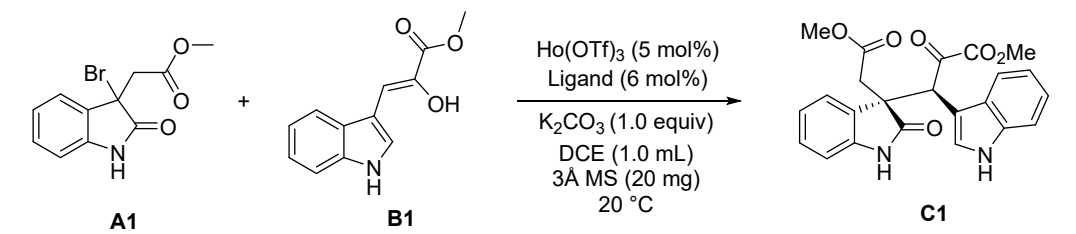
**Table S1.** Screening of metal salts



| entry <sup>a</sup> | metal salts          | yield (%) <sup>b</sup> | dr <sup>c</sup> | ee (%) <sup>c</sup> |
|--------------------|----------------------|------------------------|-----------------|---------------------|
| 1                  | Mg(OTf) <sub>2</sub> | Messy                  | -               | -                   |
| 2                  | Sc(OTf) <sub>3</sub> | Messy                  | -               | -                   |
| 3                  | Ni(OTf) <sub>2</sub> | Messy                  | -               | -                   |
| 4                  | La(OTf) <sub>3</sub> | 50                     | 69/31           | 76/70               |
| 5                  | Ce(OTf) <sub>3</sub> | 60                     | 81/19           | 78/65               |
| 6                  | Pr(OTf) <sub>3</sub> | 62                     | 86/14           | 78/55               |
| 7                  | Nd(OTf) <sub>3</sub> | 62                     | 89/11           | 85/50               |
| 8                  | Sm(OTf) <sub>3</sub> | 62                     | 79/21           | 82/57               |
| 9                  | Eu(OTf) <sub>3</sub> | 64                     | 86/14           | 88/57               |
| 10                 | Gd(OTf) <sub>3</sub> | 62                     | 80/20           | 86/51               |
| 11                 | Tb(OTf) <sub>3</sub> | 60                     | 88:12           | 86/71               |
| 12                 | Dy(OTf) <sub>3</sub> | 62                     | 90:10           | 85/70               |
| 13                 | Ho(OTf) <sub>3</sub> | 64                     | 90:10           | 88/70               |
| 14                 | Er(OTf) <sub>3</sub> | 62                     | 91:9            | 86/68               |
| 15                 | Tm(OTf) <sub>3</sub> | 52                     | 92:8            | 89/99               |
| 16                 | Yb(OTf) <sub>3</sub> | 42                     | 95:5            | 87/59               |
| 17                 | Lu(OTf) <sub>3</sub> | 40                     | 95:5            | 87/40               |
| 18 <sup>d</sup>    | Ho(OTf) <sub>3</sub> | 42                     | 62:38           | 70/13               |
| 19 <sup>e</sup>    | Ho(OTf) <sub>3</sub> | 50                     | 93:7            | 84/55               |
| 20 <sup>f</sup>    | Ho(OTf) <sub>3</sub> | 56                     | 95:5            | 84/61               |
| 21 <sup>g</sup>    | Ho(OTf) <sub>3</sub> | complex                | -               | -                   |

<sup>a</sup> The reactions were performed with **A1** (0.10 mmol), **B1** (0.1 mmol), K<sub>2</sub>CO<sub>3</sub> (0.1 mmol), 3 Å molecular sieves (20.0 mg) and metal salt/**L3-PiPr2** (1:1.2, 5 mol %) in CH<sub>2</sub>ClCH<sub>2</sub>Cl (1.0 mL) at 20 °C for 19 h. <sup>b</sup> Yield of the NMR product. <sup>c</sup> Determined by UPC<sup>2</sup> analysis on a chiral stationary phase. <sup>d</sup> Without 3 Å molecular sieves. <sup>e</sup> 4 Å molecular sieves (20.0 mg) instead of 3 Å molecular sieves. <sup>f</sup> 5 Å molecular sieves (20.0 mg) instead of 3 Å molecular sieves. <sup>g</sup> 1.8 µL H<sub>2</sub>O instead of 3 Å molecular sieves.

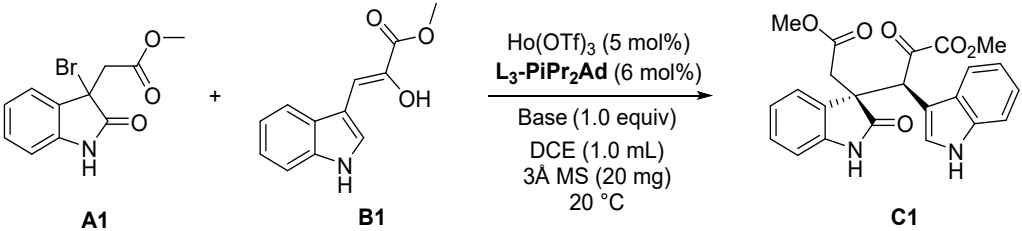
**Table S2.** Screening of ligands



| entry <sup>a</sup> | Ligand                     | yield (%) <sup>b</sup> | dr <sup>c</sup> | ee (%) <sup>c</sup> |
|--------------------|----------------------------|------------------------|-----------------|---------------------|
| 1                  | <b>L3-PiMe<sub>2</sub></b> | 48                     | 80/20           | 59/22               |
| 2                  | <b>L3-PiEt<sub>2</sub></b> | 68                     | 79/21           | 72/40               |
| 3                  | <b>L3-PiPr<sub>2</sub></b> | 64                     | 90/10           | 88/70               |
| 4                  | <b>L3-PiPr<sub>3</sub></b> | 66                     | 89:11           | 86/27               |
| 5                  | <b>L3-PiPr2Ad</b>          | 64                     | >19:1           | 92                  |
| 6                  | <b>L3-PrPr2Ad</b>          | 44                     | >19:1           | 84                  |
| 7                  | <b>L3-RaPr2Ad</b>          | 34                     | >19:1           | 35                  |

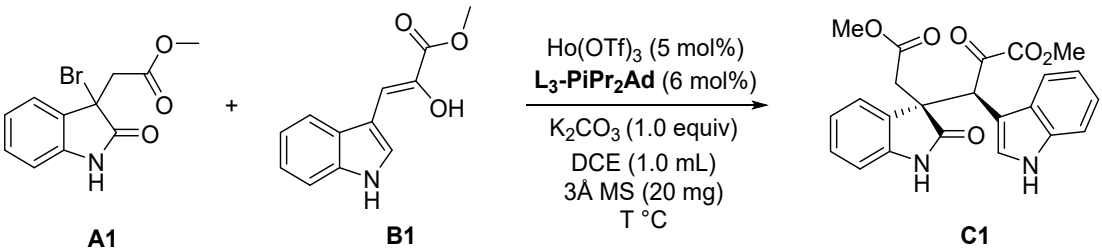
<sup>a</sup> The reactions were performed with **A1** (0.10 mmol), **B1** (0.1 mmol), K<sub>2</sub>CO<sub>3</sub> (0.1 mmol), 3 Å molecular sieves (20.0 mg) and Ho(OTf)<sub>3</sub>/**Ligand** (1:1.2, 5 mol %) in CH<sub>2</sub>ClCH<sub>2</sub>Cl (1.0 mL) at 20 °C for 19 h. <sup>b</sup> Yield of the NMR product. <sup>c</sup> Determined by UPC<sup>2</sup> analysis on a chiral stationary phase.

**Table S3.** Screening of base

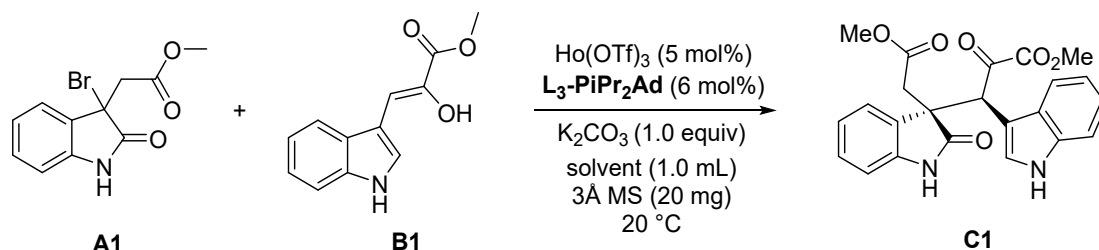
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|------------------------------------------------------------------------------------|---------------------------------|------------------------|-----------------|---------------------|
| entry <sup>a</sup>                                                                 | base                            | yield (%) <sup>b</sup> | dr <sup>c</sup> | ee (%) <sup>c</sup> |
| 1                                                                                  | KHCO <sub>3</sub>               | 46                     | >19:1           | 88                  |
| 2                                                                                  | Na <sub>2</sub> CO <sub>3</sub> | 48                     | >19:1           | 93                  |
| 3                                                                                  | K <sub>2</sub> CO <sub>3</sub>  | 64                     | >19:1           | 92                  |
| 4                                                                                  | K <sub>3</sub> PO <sub>4</sub>  | 62                     | >19:1           | 90                  |
| 5                                                                                  | Cs <sub>2</sub> CO <sub>3</sub> | complex                | -               | -                   |
| 6                                                                                  | NEt <sub>3</sub>                | complex                | -               | -                   |
| 7                                                                                  | DBU                             | complex                | -               | -                   |

<sup>a</sup> The reactions were performed with **A1** (0.10 mmol), **B1** (0.1 mmol), base (0.1 mmol), 3 Å molecular sieves (20.0 mg) and Ho(OTf)<sub>3</sub>/**L<sub>3</sub>-PiPr<sub>2</sub>Ad** (1:1.2, 5 mol %) in CH<sub>2</sub>ClCH<sub>2</sub>Cl (1.0 mL) at 20 °C for 19 h. <sup>b</sup> Yield of the NMR product. <sup>c</sup> Determined by UPC<sup>2</sup> analysis on a chiral stationary phase.

**Table S4.** Screening of temperature.

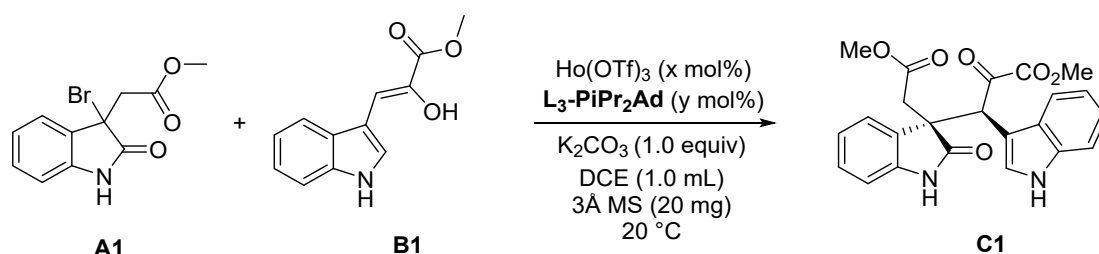
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|--------------------------------------------------------------------------------------|--------|------------------------|-----------------|---------------------|
| entry <sup>a</sup>                                                                   | T (°C) | yield (%) <sup>b</sup> | dr <sup>c</sup> | ee (%) <sup>c</sup> |
| 1 <sup>d</sup>                                                                       | -20    | trace                  | -               | -                   |
| 2 <sup>e</sup>                                                                       | 0      | 22                     | >19:1           | 93                  |
| 3 <sup>f</sup>                                                                       | 20     | 64                     | >19:1           | 92                  |
| 4 <sup>g</sup>                                                                       | 40     | 56                     | 94:6            | 87/22               |

<sup>a</sup> The reactions were performed with **A1** (0.10 mmol), **B1** (0.1 mmol), K<sub>2</sub>CO<sub>3</sub> (0.1 mmol), 3 Å molecular sieves (20.0 mg) and Ho(OTf)<sub>3</sub>/**L<sub>3</sub>-PiPr<sub>2</sub>Ad** (1:1.2, 5 mol %) in CH<sub>2</sub>ClCH<sub>2</sub>Cl (1.0 mL) at T °C. <sup>b</sup> Yield of the NMR product. <sup>c</sup> Determined by UPC<sup>2</sup> analysis on a chiral stationary phase. <sup>d</sup> Reaction time 48 h. <sup>e</sup> Reaction time 36 h. <sup>f</sup> Reaction time 19 h. <sup>g</sup> Reaction time 11 h.

**Table S5.** Screening of solvents.

| entry <sup>a</sup> | solvent           | yield (%) <sup>b</sup> | dr <sup>c</sup> | ee (%) <sup>c</sup> |
|--------------------|-------------------|------------------------|-----------------|---------------------|
| 1                  | DCM               | 46                     | >19:1           | 88                  |
| 2                  | CHCl <sub>3</sub> | 66                     | >19:1           | 73                  |
| 3                  | DCE               | 64                     | >19:1           | 92                  |
| 4                  | EA                | complex                | -               | -                   |
| 5                  | Toluene           | 16                     | >19:1           | 84                  |
| 6                  | Et <sub>2</sub> O | 20                     | >19:1           | 87                  |
| 7                  | THF               | 14                     | >19:1           | 31                  |

<sup>a</sup> The reactions were performed with **A1** (0.10 mmol), **B1** (0.1 mmol), K<sub>2</sub>CO<sub>3</sub> (0.1 mmol), 3 Å molecular sieves (20.0 mg) and Ho(OTf)<sub>3</sub>/ **L3-PiPr<sub>2</sub>Ad** (1:1.2, 5 mol %) in solvent (1.0 mL) at 20 °C for 19 h. <sup>b</sup> Yield of the NMR product. <sup>c</sup> Determined by UPC<sup>2</sup> analysis on a chiral stationary phase.

**Table S6.** Ratio of Ho(OTf)<sub>3</sub> and **L3-PiPr<sub>2</sub>Ad**.

| entry <sup>a</sup> | x:y   | yield (%) <sup>b</sup> | dr <sup>c</sup> | ee (%) <sup>c</sup> |
|--------------------|-------|------------------------|-----------------|---------------------|
| 1                  | 7.5:5 | 68                     | >19:1           | 70                  |
| 2                  | 6:5   | 64                     | >19:1           | 70                  |
| 3                  | 5:5   | 64                     | >19:1           | 90                  |
| 4                  | 5:6   | 64                     | >19:1           | 92                  |
| 5                  | 5:7.5 | 64                     | >19:1           | 92                  |
| 6                  | 5:10  | 69                     | >19:1           | 93                  |

<sup>a</sup> The reactions were performed with **A1** (0.10 mmol), **B1** (0.1 mmol), K<sub>2</sub>CO<sub>3</sub> (0.1 mmol), 3Å molecular sieves (20.0 mg) and Ho(OTf)<sub>3</sub>/**L3-PiPr<sub>2</sub>Ad** (x:y) in CH<sub>2</sub>ClCH<sub>2</sub>Cl (1.0 mL) at 20 °C for 19 h. <sup>b</sup> Yield of the NMR product. <sup>c</sup> Determined by UPC<sup>2</sup> analysis on a chiral stationary phase.

**Table S7.** Screening of ratio of substrate.

| entry <sup>a</sup> | x:y      | yield (%) <sup>b</sup> | dr <sup>c</sup> | ee (%) <sup>c</sup> |
|--------------------|----------|------------------------|-----------------|---------------------|
| 1 <sup>d</sup>     | 0.2:0.1  | 95                     | >19:1           | 91                  |
| 2 <sup>d</sup>     | 0.15:1   | 95                     | >19:1           | 92                  |
| 3                  | 0.1:0.1  | 64                     | >19:1           | 91                  |
| 4                  | 0.1:0.15 | 64                     | >19:1           | 90                  |
| 5                  | 0.1:0.2  | 62                     | >19:1           | 91                  |

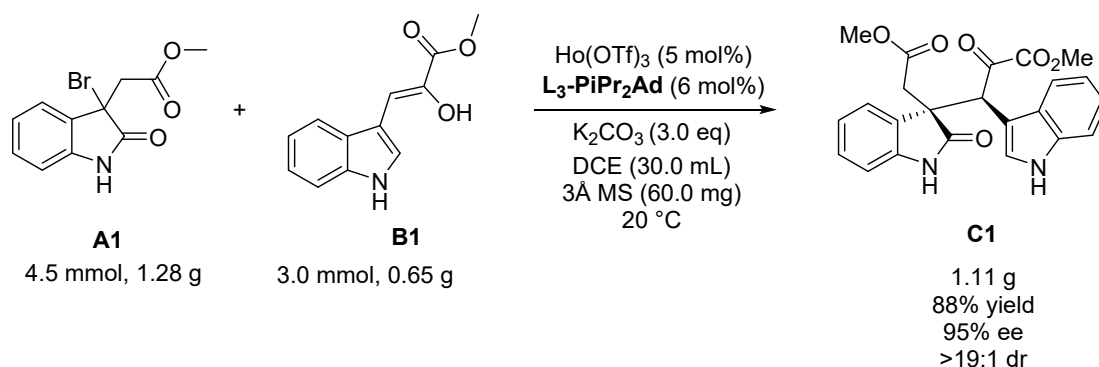
<sup>a</sup> The reactions were performed with **A1** (x mmol), **B1** (y mmol), K<sub>2</sub>CO<sub>3</sub> (0.1 mmol), 3 Å molecular sieves (20.0 mg) and Ho(OTf)<sub>3</sub>/ **L3-PiPr<sub>2</sub>Ad** (1:1.2, 5 mol %) in DCE (1.0 mL) at 20 °C for 19 h. <sup>b</sup> Yield of the NMR product. <sup>c</sup> Determined by UPC<sup>2</sup> analysis on a chiral stationary phase. <sup>d</sup> Reaction time 11 h.

**Table S8.** Screening of the amount of base.

| entry <sup>a</sup> | the amount of K <sub>2</sub> CO <sub>3</sub><br>(x mmol) | yield (%) <sup>b</sup> | dr <sup>c</sup> | ee (%) <sup>c</sup> |
|--------------------|----------------------------------------------------------|------------------------|-----------------|---------------------|
| 1                  | 0.1                                                      | 95                     | >19:1           | 92                  |
| 2                  | 0.2                                                      | 95                     | >19:1           | 94                  |
| 3                  | 0.3                                                      | 95                     | >19:1           | 96                  |

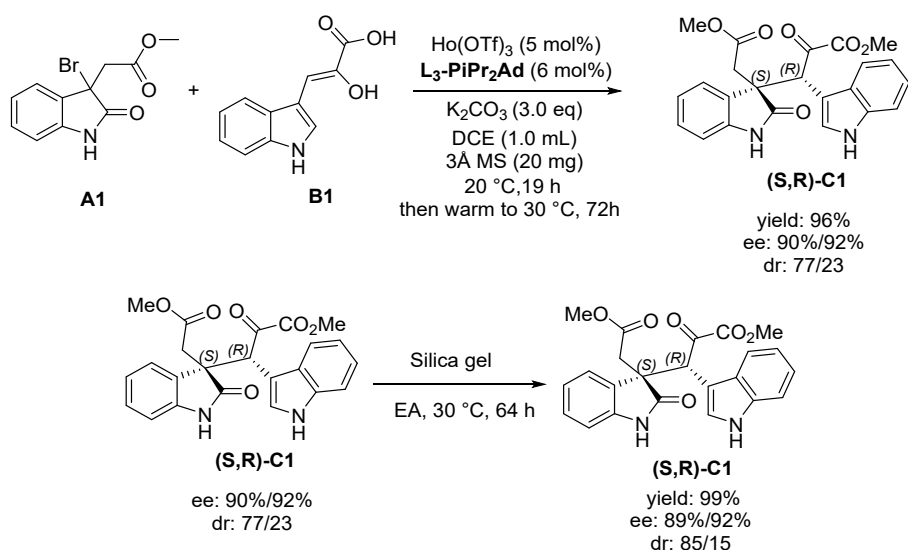
<sup>a</sup> The reactions were performed with **A1** (0.15 mmol), **B1** (0.1 mmol), K<sub>2</sub>CO<sub>3</sub> (x mmol), 3 Å molecular sieves (20.0 mg) and Ho(OTf)<sub>3</sub>/ **L3-PiPr<sub>2</sub>Ad** (1:1.2, 5 mol %) in DCE (1.0 mL) at 20 °C for 11 h. <sup>b</sup> Yield of the NMR product. <sup>c</sup> Determined by UPC<sup>2</sup> analysis on a chiral stationary phase.

#### 4 Gram-scale synthesis of C1



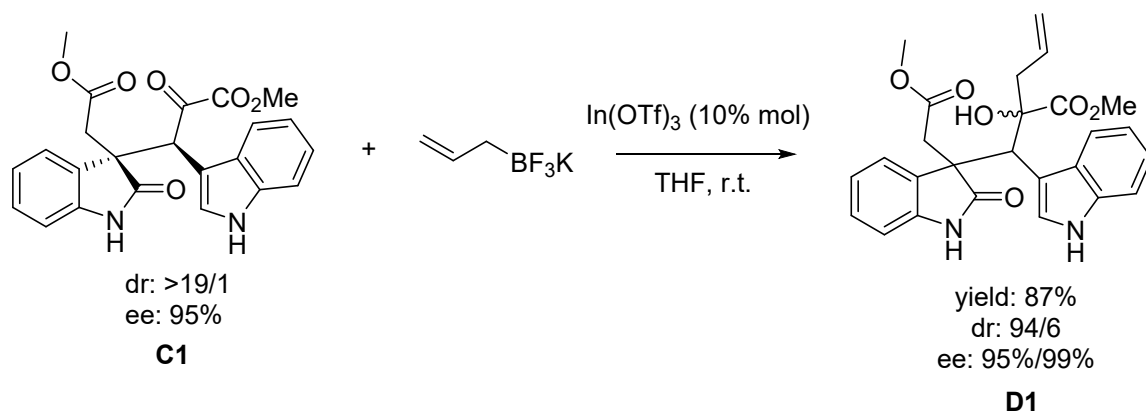
An oven-dried 100 mL flask was charged with *N,N*-dioxide L<sub>3</sub>-PiPr<sub>2</sub>Ad (165.1 mg, 0.18 mmol), Ho(OTf)<sub>3</sub> (91.8 mg, 0.15 mmol), and DCM (7.5 mL) under N<sub>2</sub> atmosphere. After the mixture was stirred at 35 °C for 30 min, the solvent was removed in vacuo. 3-Bromo-3-substituted oxindoles **A1** (1.28 g, 4.5 mmol), methyl (Z)-2-hydroxy-3-(1H-indol-3-yl)acrylate **B1** (0.65 g, 3 mmol), K<sub>2</sub>CO<sub>3</sub> (1.24 g, 9 mmol, 3.0 equiv.), 3Å MS. (600.0 mg), and 30.0 mL DCE were added subsequently in the glovebox. The reaction was performed at 20 °C for 16 hours. The reaction mixture was directly subjected to flash column chromatography on silica gel at -35 °C (eluent: petroleum ether/ethyl acetate = 2:1 to 1:1, v/v) to afford the corresponding products **C1**. By the way, the product suffered from somewhat epimerization after the solvent was removed in vacuo.

#### 5 Synthetic transformations



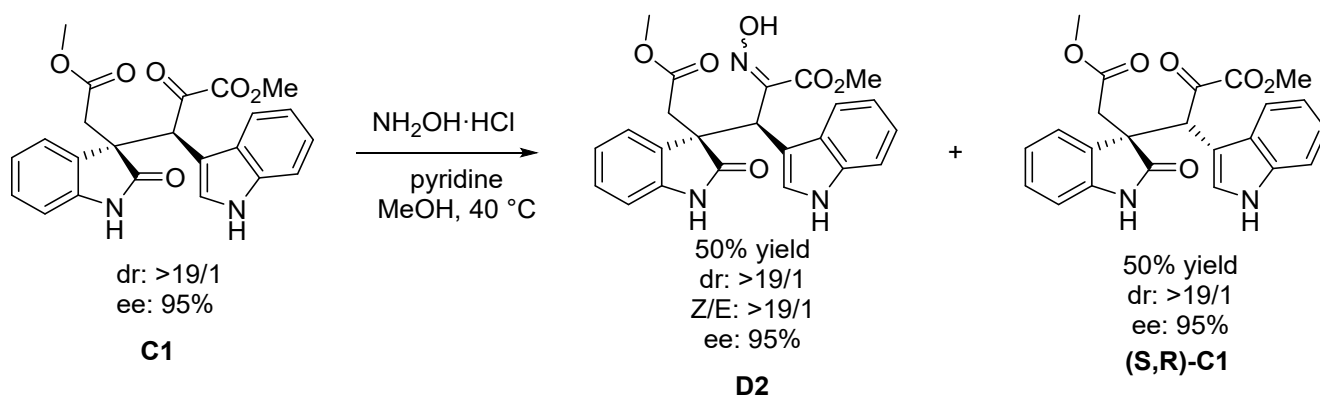
An oven-dried test tube was charged with *N,N*-dioxide L<sub>3</sub>-PiPr<sub>2</sub>Ad (5.5 mg, 0.006 mmol), Ho(OTf)<sub>3</sub> (3.1 mg, 0.005 mmol), and 0.5 mL DCM under N<sub>2</sub> atmosphere. After the mixture was stirred at 35 °C for 30 min, the solvent was removed in vacuo. 3-Bromo-3-substituted oxindoles **A1** (42.6 mg, 0.15 mmol), methyl (Z)-2-hydroxy-3-(1H-indol-3-yl)acrylate **B1** (21.7 mg, 0.1 mmol), K<sub>2</sub>CO<sub>3</sub> (41.4 mg, 0.3 mmol, 3.0 equiv.), 3Å MS. (20 mg), and 1.0 mL DCE were added in the glovebox. The reaction was performed at 20 °C. After 19 h, the reaction was performed at 30 °C for 72 h. The reaction mixture was directly subjected to flash column chromatography on silica gel at -35 °C (eluent: petroleum ether/ethyl acetate = 2:1 to 1:1, v/v) to afford the corresponding products **(S,R)-C1**. An oven-dried test tube was charged with **(S,R)-C1** (0.05 mmol, 21.1 mg), 0.3 g Silica gel and 1 mL EA. The reaction was performed at 30 °C for 64 h. The reaction mixture was directly subjected to flash filtration to afford the **(S,R)-C1** in 99% yield. The dr value and ee value were determined by UPC<sup>2</sup> analysis using a chiral stationary phase.

##### 5.1 Procedure for the synthesis of D1

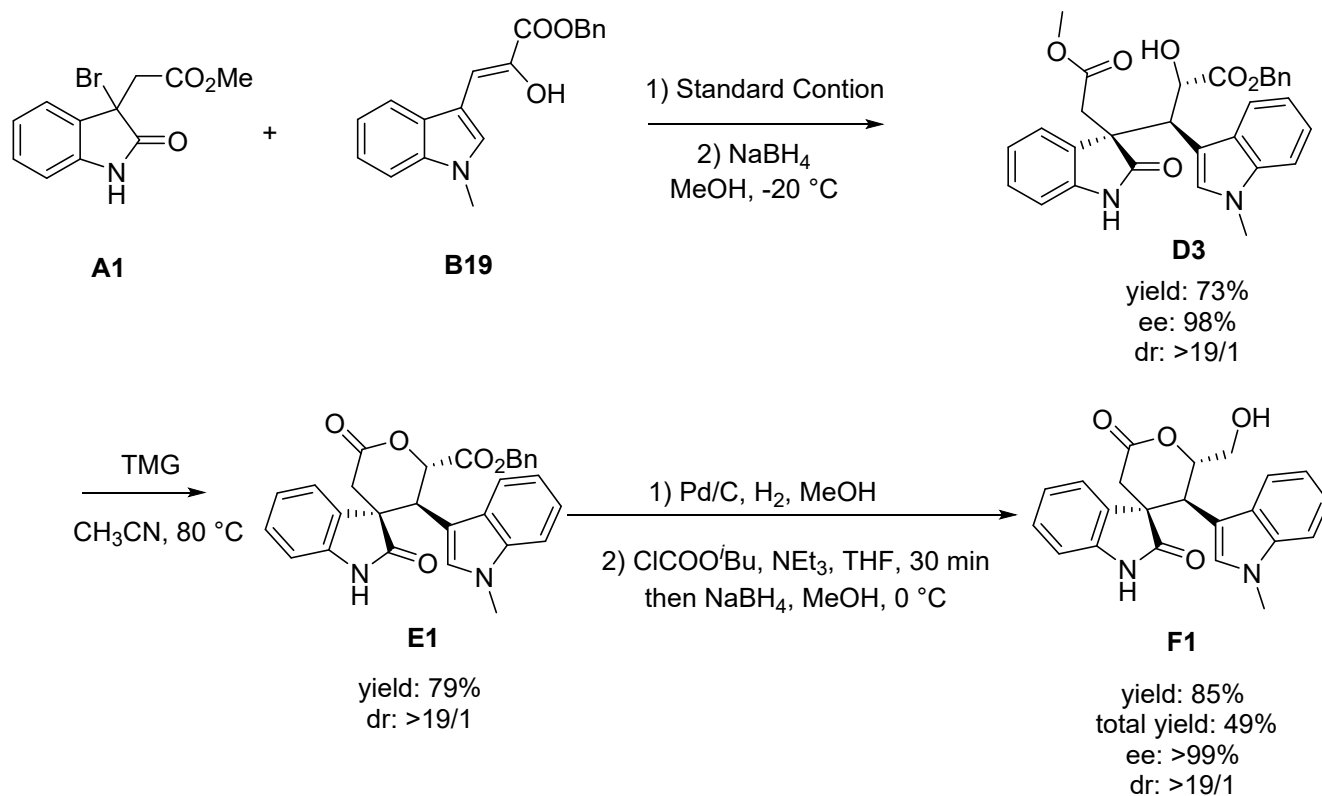


An over-dried test tube was charged with  $\text{In(OTf)}_3$  (0.01 mmol, 5.6 mg), **C1** (0.1 mmol, 42.1 mg) and 1 mL THF in the glovebox. The mixture was stirred at room temperature for 30 min. Potassium allyltrifluoroborate was added (0.2 mmol, 29.6 mg) to the test tube. The reaction was performed at room temperature for 30 min. The reaction mixture was directly subjected to flash column chromatography on silica gel (eluent: petroleum ether/ethyl acetate = 1:1 to 1:2, v/v) to afford the corresponding products **D1** in 87% yield. The dr value was determined by NMR. The ee value was determined by UPC<sup>2</sup> analysis using a chiral stationary phase.

## 5.2 Procedure for the synthesis of D2

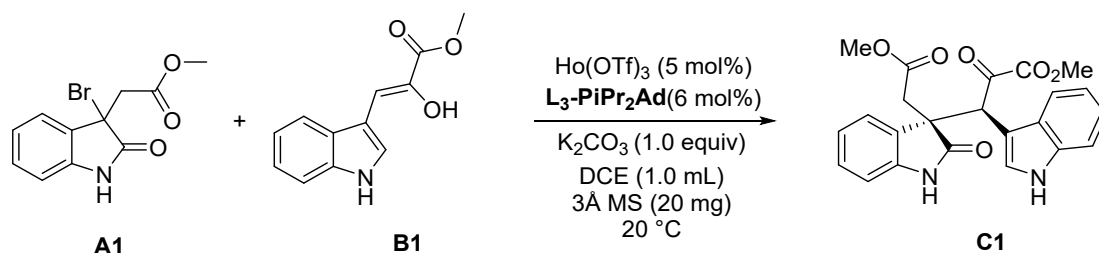


An over-dried test tube was charged with  $\text{NH}_2\text{OH}\cdot\text{HCl}$  (0.12 mmol, 8.4 mg), pyridine (0.11 mmol, 8.9  $\mu\text{L}$ ) and 1.0 mL MeOH. The mixture was stirred at 40 °C for 0.5 h. **C1** (0.1 mmol, 42.1 mg) was added to the test tube. The reaction was performed at the 40 °C for 36 h. The reaction mixture was directly subjected to flash column chromatography on silica gel (eluent: petroleum ether/ethyl acetate = 1:1 to 2:3, v/v) to afford the corresponding products **D1** in 50% yield. The dr value and Z/E value were determined by NMR. The ee value was determined by UPC<sup>2</sup> analysis using a chiral stationary phase.



An oven-dried test tube was charged with *N,N'*-dioxide **L<sub>3</sub>-PiPr<sub>2</sub>Ad** (5.5 mg, 0.006 mmol), Ho(OTf)<sub>3</sub> (3.1 mg, 0.005 mmol), and 0.5 mL DCM under N<sub>2</sub> atmosphere. After the mixture was stirred at 35 °C for 30 min, the solvent was removed in vacuo. 3-Bromo-3-substituted oxindoles **A1** (42.6 mg, 0.15 mmol), benzyl (Z)-2-hydroxy-3-(1-methyl-1H-indol-3-yl)acrylate **B19** (21.7 mg, 0.1 mmol), K<sub>2</sub>CO<sub>3</sub> (41.4 mg, 0.3 mmol, 3.0 equiv.), 3Å MS. (20 mg), and 1.0 mL DCE were added subsequently in the glovebox. The reaction was performed at 20 °C. After 15 h, The reaction mixture was directly subjected to flash filtration, the solvent was removed in vacuo. The mixture was dissolved in 2.0 mL MeOH at -20°C. NaBH<sub>4</sub> (11.4 mg, 0.3 mmol) was added. The reaction was performed at -20 °C for 30 minutes. The reaction mixture was directly subjected to flash column chromatography on silica gel (eluent: petroleum ether/ethyl acetate = 3:2 to 1:1, v/v) to afford the corresponding products **D3** in 73% yield. **D3** (51.2 mg, 0.1 mmol), TMG (12.55 μL, 0.1 mmol), and 1.0 mL CH<sub>3</sub>CN were added to a test tube. The reaction was performed at 80 °C for 13 h. The reaction mixture was directly subjected to flash column chromatography on silica gel (eluent: petroleum ether/ethyl acetate = 3:2 to 1:1 v/v,) to afford the corresponding products **E1** in 79% yield. An oven-dried test tube was charged with **E1** (48.1 mg, 0.1 mmol), Pd/C (4.0 mg), and 2 mL MeOH under H<sub>2</sub> atmosphere. After 16h, the product was filtered with coarse silica gel, the solvent was removed in vacuo. the mixture was dissolved in 2.0 mL THF. NEt<sub>3</sub> (15.3 uL, 0.11 mmol) and CICOOBu (16.5 uL, 0.12 mmol) were added under N<sub>2</sub> atmosphere. The reaction was performed at room temperature for 30 minutes, and then 1.0 mL MeOH and NaBH<sub>4</sub> (11.4 mg, 0.3 mmol) were added at 0 °C. After for 30 minutes, the reaction mixture was directly subjected to flash column chromatography on silica gel (eluent: petroleum ether/ethyl acetate = 1:1 to 1:2, v/v) to afford the corresponding products **F1** in 85% yield. The dr value was determined by NMR and UPC<sup>2</sup> analysis using a chiral stationary phase. The ee value was determined by UPC<sup>2</sup> analysis using a chiral stationary phase.

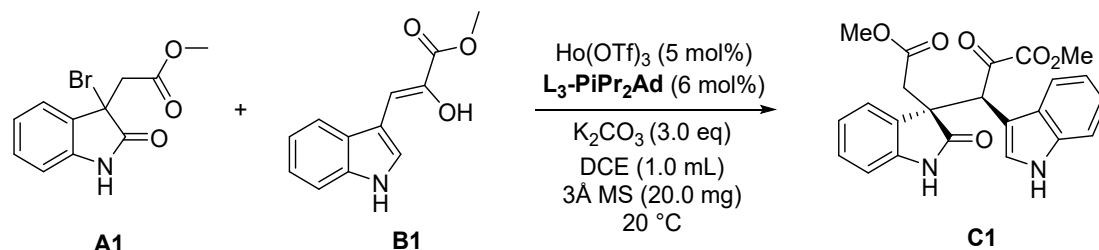
## 6 Control experiments



| entry <sup>a</sup> | Variations                                                            | yield (%) <sup>b</sup> | dr <sup>c</sup> | ee (%) <sup>c</sup> |
|--------------------|-----------------------------------------------------------------------|------------------------|-----------------|---------------------|
| 1                  | -                                                                     | 64                     | >19:1           | 92                  |
| 2                  | Without Ho(OTf) <sub>3</sub> and L <sub>3</sub> -PiPr <sub>2</sub> Ad | trace                  | -               | -                   |
| 3                  | Without L <sub>3</sub> -PiPr <sub>2</sub> Ad                          | 62                     | 9/1             | -                   |

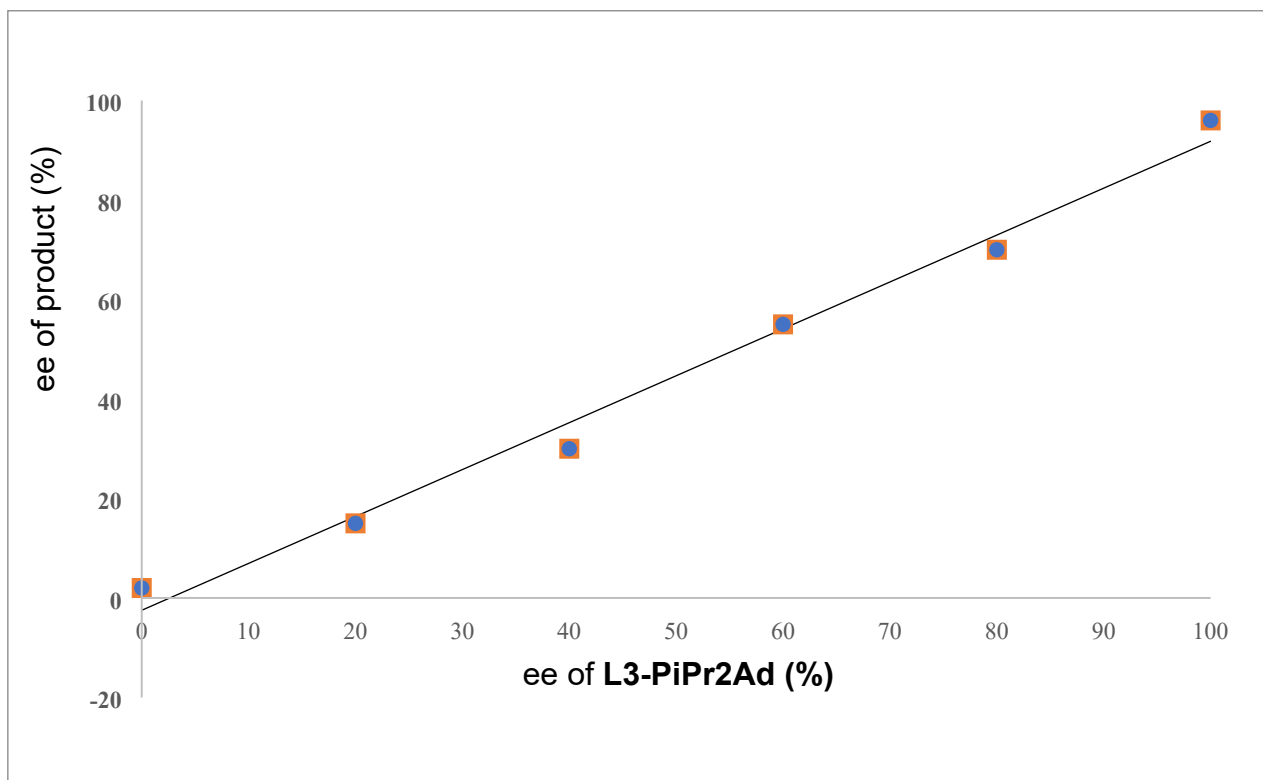
<sup>a</sup> The reactions were performed with **A1** (0.10 mmol), **B1** (0.1 mmol), K<sub>2</sub>CO<sub>3</sub> (0.1 mmol), 3Å molecular sieves (20.0 mg) and Ho(OTf)<sub>3</sub>/L<sub>3</sub>-PiPr<sub>2</sub>Ad (1:1.2, 5 mol %) in CH<sub>2</sub>ClCH<sub>2</sub>Cl (1.0 mL) at 20 °C for 19 h. <sup>b</sup> Yield of the NMR product. <sup>c</sup> Determined by UPC<sup>2</sup> analysis on a chiral stationary phase.

## 7 The relationship between ee of the ligand and ee of the product C1

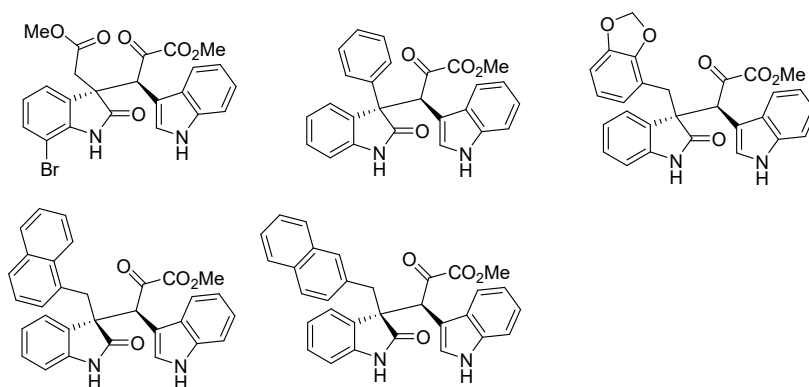


| entry <sup>a</sup> | ee of ligand | yield (%) <sup>b</sup> | dr <sup>c</sup> | ee (%) <sup>c</sup> |
|--------------------|--------------|------------------------|-----------------|---------------------|
| 1                  | 0            | 94                     | >19:1           | 2                   |
| 2                  | 20           | 95                     | >19:1           | 15                  |
| 3                  | 40           | 96                     | >19:1           | 30                  |
| 4                  | 60           | 94                     | >19:1           | 55                  |
| 5                  | 80           | 95                     | >19:1           | 70                  |
| 6                  | 100          | 95                     | >19:1           | 96                  |

<sup>a</sup> The reactions were performed with **A1** (0.15 mmol), **B1** (0.1 mmol), K<sub>2</sub>CO<sub>3</sub> (0.3 mmol), 3Å molecular sieves (20.0 mg) and Ho(OTf)<sub>3</sub>/L<sub>3</sub>-PiPr<sub>2</sub>Ad (1:1.2, 5 mol %) in CH<sub>2</sub>ClCH<sub>2</sub>Cl (1.0 mL) at 20 °C for 11 h. <sup>b</sup> Yield of the NMR product. <sup>c</sup> Determined by UPC<sup>2</sup> analysis on a chiral stationary phase.

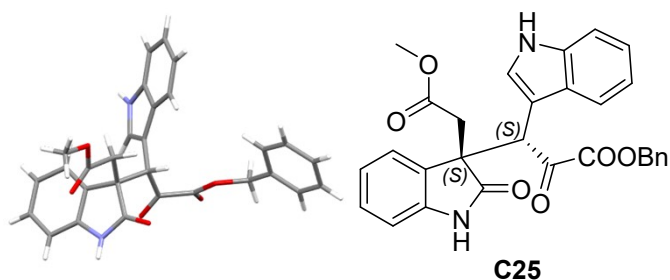


#### 8 Unsuccessful substrate scopes



## 9 Determination of absolute configuration of compound **C25**

Crystals suitable for the X-ray crystal structure analysis were obtained from a solution of compound **C25** in  $\text{CHCl}_3$  (ca. 5 mL) at room temperature. The yellow crystal in rod-shape, with approximate dimensions of  $0.194 \times 0.244 \times 0.577 \text{ mm}^3$ , was selected and mounted for the single-crystal X-ray diffraction. The data set was collected by Bruker D8 Venture Photon II diffractometer at 173(2)K equipped with micro-focus Cu radiation source ( $K\alpha = 1.54178\text{\AA}$ ). Applied with face-indexed numerical absorption correction, the structure solution was solved and refinement was processed by SHELXTL (version 6.14) and OLEX 2.3 program package.<sup>[4]</sup> The structure was analyzed by ADDSYM routine implemented in PLATON suite and no higher symmetry was suggested.<sup>[5]</sup>



Crystallographic Data for  $C_{30}H_{25}Cl_3N_2O_6$

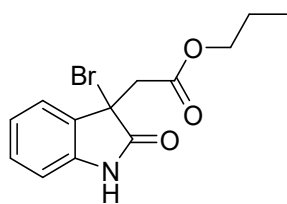
|                                                   |                          |
|---------------------------------------------------|--------------------------|
| Formula                                           | $C_{30}H_{25}Cl_3N_2O_6$ |
| Formula mass (amu)                                | 615.87                   |
| Space group                                       | P 21 21 21               |
| $a$ (Å)                                           | 9.4395(2)                |
| $b$ (Å)                                           | 17.2117(4)               |
| $c$ (Å)                                           | 18.1895(4)               |
| $\alpha$ (deg)                                    | 90                       |
| $\beta$ (deg)                                     | 90                       |
| $\gamma$ (deg)                                    | 90                       |
| $V$ (Å <sup>3</sup> )                             | 2955.25(11)              |
| $Z$                                               | 4                        |
| $\lambda$ (Å)                                     | 1.54178                  |
| $T$ (K)                                           | 173                      |
| $\rho_{\text{calcd}}$ (g cm <sup>-3</sup> )       | 1.384                    |
| $\mu$ (mm <sup>-1</sup> )                         | 3.195                    |
| Transmission factors                              | 0.475-0.944              |
| $2\theta_{\text{max}}$ (deg)                      | 68.241                   |
| No. of unique data, including $F_o^2 < 0$         | 5364                     |
| No. of unique data, with $F_o^2 > 2\sigma(F_o^2)$ | 5204                     |
| No. of variables                                  | 379                      |
| $R(F)$ for $F_o^2 > 2\sigma(F_o^2)$ <sup>a</sup>  | 0.0354                   |
| $R_w(F_o^2)$ <sup>b</sup>                         | 0.0916                   |
| Goodness of fit                                   | 1.040                    |

<sup>a</sup>  $R(F) = \sum ||F_o| - |F_c|| / \sum |F_o|$ .

<sup>b</sup>  $R_w(F_o^2) = [\sum [w(F_o^2 - F_c^2)^2] / \sum wF_o^4]^{1/2}$ ;  $w^{-1} = [\sigma^2(F_o^2) + (Ap)^2 + Bp]$ , where  $p = [\max(F_o^2, 0) + 2F_c^2] / 3$ .

## 10 Characterization of the products

### Propyl 2-(3-bromo-2-oxoindolin-3-yl)acetate (A3)



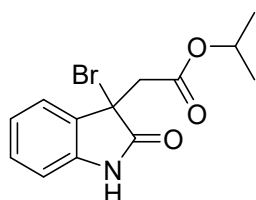
**A3**

Yellow solid.

$^1\text{H}$  NMR (600 MHz, Chloroform-*d*)  $\delta$  9.11 (s, 1H), 7.37 (s, 1H), 7.06 (s, 1H), 6.97 – 6.90 (m, 2H), 3.86 (t,  $J$  = 6.0, 2H), 3.62 (dd,  $J$  = 48.0, 18.0 Hz, 2H), 1.43 (s, 2H), 0.77 (s, 3H).

$^{13}\text{C}$  NMR (151 MHz,  $\text{CDCl}_3$ )  $\delta$  176.4, 168.2, 140.4, 130.6, 129.8, 124.1, 123.3, 111.0, 67.0, 51.4, 43.4, 21.7, 10.3.

### Isopropyl 2-(3-bromo-2-oxoindolin-3-yl)acetate (A4)



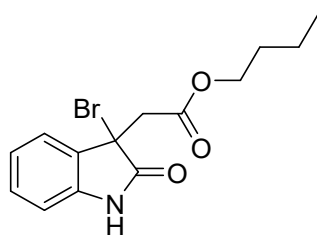
**A4**

Yellow solid.

$^1\text{H}$  NMR (600 MHz, Chloroform-*d*)  $\delta$  8.92 (s, 1H), 7.36 (d,  $J$  = 7.6 Hz, 1H), 7.27 – 7.25 (m, 1H), 7.05 (t,  $J$  = 7.5 Hz, 1H), 6.93 (d,  $J$  = 7.8 Hz, 1H), 4.76 (p,  $J$  = 6.2 Hz, 1H), 3.56 (dd,  $J$  = 64.0, 18.0 Hz, 2H), 1.03 (d,  $J$  = 6.2 Hz, 3H), 0.92 (d,  $J$  = 6.2 Hz, 3H).

$^{13}\text{C}$  NMR (151 MHz,  $\text{CDCl}_3$ )  $\delta$  176.4, 167.5, 140.4, 130.6, 129.9, 124.1, 123.3, 110.9, 69.2, 51.4, 43.8, 21.5, 21.3.

### Butyl 2-(3-bromo-2-oxoindolin-3-yl)acetate (A5)



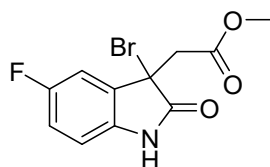
**A5**

Yellow solid.

$^1\text{H}$  NMR (400 MHz, Chloroform-*d*)  $\delta$  8.54 (s, 1H), 7.37 (d,  $J$  = 7.5 Hz, 1H), 7.31 – 7.23 (m, 1H), 7.07 (t,  $J$  = 7.6 Hz, 1H), 6.92 (d,  $J$  = 7.8 Hz, 1H), 4.05 – 3.78 (m, 2H), 3.61 (dd,  $J$  = 48.0, 16.0 Hz, 2H), 1.45 – 1.34 (m, 2H), 1.22 – 1.14 (m, 2H), 0.82 (t,  $J$  = 7.3 Hz, 3H).

$^{13}\text{C}$  NMR (101 MHz,  $\text{CDCl}_3$ )  $\delta$  175.9, 168.3, 140.3, 130.6, 129.9, 124.1, 123.4, 110.8, 65.3, 51.2, 43.5, 30.4, 19.0, 13.7.

### Methyl 2-(3-bromo-5-fluoro-2-oxoindolin-3-yl)acetate (A6)



**A6**

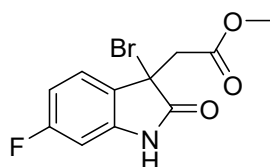
Yellow solid.

$^1\text{H}$  NMR (400 MHz, Acetonitrile- $d_3$ )  $\delta$  8.71 (s, 1H), 7.25 (dd,  $J$  = 8.2, 2.7 Hz, 1H), 7.05 (td,  $J$  = 9.1, 2.6 Hz, 1H), 6.92 (dd,  $J$  = 8.6, 4.3 Hz, 1H), 3.63 (d,  $J$  = 17.2 Hz, 1H), 3.53 – 3.49 (m, 4H).

$^{13}\text{C}$  NMR (101 MHz, Acetonitrile- $d_3$ )  $\delta$  175.65, 169.72, 159.71 (d,  $J$  = 239.0 Hz), 138.00 (d,  $J$  = 2.1 Hz), 132.32 (d,  $J$  = 9.1 Hz), 117.64 (d,  $J$  = 23.9 Hz), 112.67 (d,  $J$  = 25.5 Hz), 112.31 (d,  $J$  = 8.0 Hz).

$^{19}\text{F}$  NMR (377 MHz,  $\text{CD}_3\text{CN}$ )  $\delta$  -121.8.

**Methyl 2-(3-bromo-6-fluoro-2-oxoindolin-3-yl)acetate (A7)**



**A7**

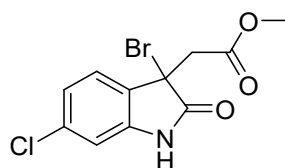
Yellow solid.

$^1\text{H}$  NMR (400 MHz, Acetonitrile- $d_3$ )  $\delta$  8.81 (s, 1H), 7.43 (dd,  $J$  = 8.4, 5.4 Hz, 1H), 6.84 – 6.68 (m, 2H), 3.65 (d,  $J$  = 17.1 Hz, 1H), 3.513 – 3.49 (m,  $J$  = 15.9 Hz, 4H).

$^{13}\text{C}$  NMR (101 MHz, Acetonitrile- $d_3$ )  $\delta$  175.87, 169.73, 164.71 (d,  $J$  = 246.2 Hz), 143.59 (d,  $J$  = 12.5 Hz), 126.70 (d,  $J$  = 10.1 Hz), 126.69, 109.89 (d,  $J$  = 23.3 Hz), 99.74 (d,  $J$  = 27.8 Hz), 52.55, 51.78, 43.14.

$^{19}\text{F}$  NMR (377 MHz,  $\text{CD}_3\text{CN}$ )  $\delta$  -110.3.

**Methyl 2-(3-bromo-6-chloro-2-oxoindolin-3-yl)acetate (A8)**



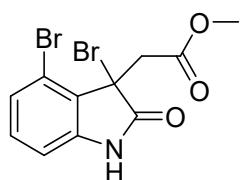
**A8**

Yellow solid.

$^1\text{H}$  NMR (400 MHz, Chloroform- $d$ )  $\delta$  8.56 (s, 1H), 7.28 (d,  $J$  = 8.1 Hz, 1H), 7.04 (dd,  $J$  = 8.1, 1.9 Hz, 1H), 6.94 (d,  $J$  = 1.9 Hz, 1H), 3.65 (d,  $J$  = 17.1 Hz, 1H), 3.56 – 3.51 (m, 4H).

$^{13}\text{C}$  NMR (101 MHz,  $\text{CDCl}_3$ )  $\delta$  175.8, 168.8, 141.4, 136.4, 128.2, 125.1, 123.5, 111.6, 52.4, 50.1, 43.1.

**Methyl 2-(3,4-dibromo-2-oxoindolin-3-yl)acetate (A9)**



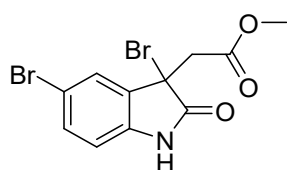
**A9**

Yellow solid.

$^1\text{H}$  NMR (600 MHz, Chloroform-*d*)  $\delta$  8.60 (s, 1H), 7.18 (d,  $J$  = 8.1 Hz, 1H), 7.14 (t,  $J$  = 7.9 Hz, 1H), 6.89 (d,  $J$  = 7.6 Hz, 1H), 3.90 (dd,  $J$  = 364.9, 17.4 Hz, 2H), 3.56 (s, 3H).

$^{13}\text{C}$  NMR (151 MHz,  $\text{CDCl}_3$ )  $\delta$  175.2, 169.2, 142.4, 131.7, 127.6, 127.4, 119.9, 109.9, 52.3, 51.8, 41.3.

**Methyl 2-(3,5-dibromo-2-oxoindolin-3-yl)acetate (A10)**



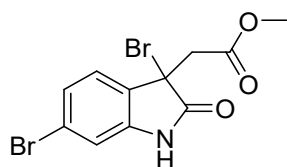
**A10**

Yellow solid.

$^1\text{H}$  NMR (600 MHz, Chloroform-*d*)  $\delta$  8.16 (s, 1H), 7.47 (d,  $J$  = 2.1 Hz, 1H), 7.40 (dd,  $J$  = 8.2, 2.0 Hz, 1H), 6.81 (d,  $J$  = 8.3 Hz, 1H), 3.65 (d,  $J$  = 17.3 Hz, 1H), 3.57 (s, 3H), 3.51 (d,  $J$  = 17.3 Hz, 1H).

$^{13}\text{C}$  NMR (151 MHz,  $\text{CDCl}_3$ )  $\delta$  175.1, 168.8, 139.2, 133.4, 131.9, 127.3, 115.8, 112.3, 52.4, 49.9, 43.0.

**Methyl 2-(3,6-dibromo-2-oxoindolin-3-yl)acetate (A11)**

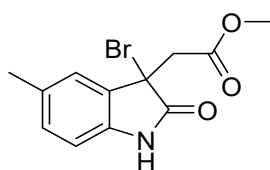


**A11**

$^1\text{H}$  NMR (600 MHz, Chloroform-*d*)  $\delta$  8.90 (s, 1H), 7.25 – 7.17 (m, 2H), 7.11 (d,  $J$  = 1.7 Hz, 1H), 3.65 (d,  $J$  = 17.1 Hz, 1H), 3.59 – 3.50 (m, 4H).

$^{13}\text{C}$  NMR (151 MHz,  $\text{CDCl}_3$ )  $\delta$  175.9, 168.8, 141.6, 128.8, 126.4, 125.3, 124.3, 114.5, 52.4, 50.2, 43.0.

**Methyl 2-(3-bromo-5-methyl-2-oxoindolin-3-yl)acetate (A12)**



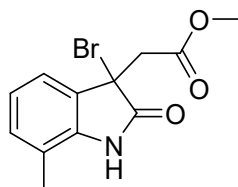
**A12**

Yellow solid.

$^1\text{H}$  NMR (600 MHz, Chloroform-*d*)  $\delta$  8.72 (s, 1H), 7.17 (d,  $J$  = 1.5 Hz, 1H), 7.06 (dd,  $J$  = 8.0, 1.6 Hz, 1H), 6.81 (d,  $J$  = 7.9 Hz, 1H), 3.65 (d,  $J$  = 17.0 Hz, 1H), 3.56 -3.53 (m, 4H), 2.31 (s, 3H).

$^{13}\text{C}$  NMR (151 MHz,  $\text{CDCl}_3$ )  $\delta$  176.1, 168.8, 137.9, 133.0, 131.1, 129.7, 124.7, 110.7, 52.3, 51.5, 43.2, 21.2.

**Methyl 2-(3-bromo-7-methyl-2-oxoindolin-3-yl)acetate (A13)**



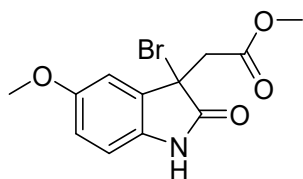
**A13**

Yellow solid.

$^1\text{H}$  NMR (600 MHz, Chloroform-*d*)  $\delta$  9.37 (s, 1H), 7.20 (d,  $J$  = 7.5 Hz, 1H), 7.09 (d,  $J$  = 7.7 Hz, 1H), 6.97 (t,  $J$  = 7.6 Hz, 1H), 3.61 (dd,  $J$  = 60.0, 18.0 Hz 2H), 3.51 (s, 3H), 2.31 (s, 3H).

$^{13}\text{C}$  NMR (151 MHz,  $\text{CDCl}_3$ )  $\delta$  176.7, 168.8, 139.2, 132.0, 129.3, 123.2, 121.4, 120.5, 52.2, 51.9, 43.1, 16.5.

**methyl 2-(3-bromo-5-methoxy-2-oxoindolin-3-yl)acetate (A14)**



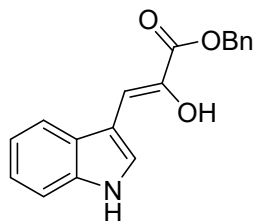
**A14**

Yellow solid.

$^1\text{H}$  NMR (400 MHz, Chloroform-*d*)  $\delta$  7.98 (s, 1H), 7.02 – 6.91 (m, 1H), 6.81 (t,  $J$  = 1.6 Hz, 2H), 3.79 (s, 3H), 3.64 (d,  $J$  = 17.0 Hz, 1H), 3.55 – 3.49 (d, 4H).

$^{13}\text{C}$  NMR (101 MHz,  $\text{CDCl}_3$ )  $\delta$  175.5, 168.8, 156.3, 133.4, 130.9, 115.4, 111.3, 110.9, 55.9, 52.3, 51.3, 43.2.

**Benzyl (Z)-2-hydroxy-3-(1H-indol-3-yl)acrylate (B2)**



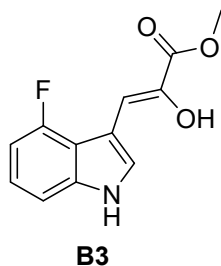
**B2**

Red solid.

$^1\text{H}$  NMR (600 MHz, Chloroform-*d*)  $\delta$  8.41 (s, 1H), 7.98 (d,  $J$  = 2.6 Hz, 1H), 7.78 (d,  $J$  = 7.8 Hz, 1H), 7.46 (d,  $J$  = 7.1 Hz, 2H), 7.44 – 7.37 (m, 4H), 7.27 – 7.23 (m, 1H), 7.20 (t,  $J$  = 7.5 Hz, 1H), 7.01 (s, 1H), 6.25 (s, 1H), 5.36 (s, 2H).

$^{13}\text{C}$  NMR (151 MHz,  $\text{CDCl}_3$ )  $\delta$  166.1, 136.7, 135.7, 135.6, 128.8, 128.6, 128.4, 127.6, 126.9, 122.9, 120.6, 118.8, 111.4, 110.8, 104.5, 67.8.

**Methyl (Z)-3-(4-fluoro-1H-indol-3-yl)-2-hydroxyacrylate (B3)**



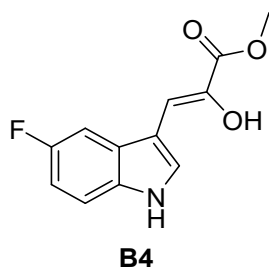
Yellow solid.

$^1\text{H}$  NMR (600 MHz, Chloroform- $d$ )  $\delta$  8.50 (s, 1H), 7.98 (d,  $J$  = 2.3 Hz, 1H), 7.21 (s, 1H), 7.18 – 7.11 (m, 2H), 6.84 (dd,  $J$  = 11.2, 7.6 Hz, 1H), 3.92 (s, 3H).

$^{13}\text{C}$  NMR (151 MHz, Chloroform- $d$ )  $\delta$  166.77, 157.60 (d,  $J$  = 247.3 Hz), 138.15 (d,  $J$  = 11.0 Hz), 136.91, 127.64, 123.29 (d,  $J$  = 8.5 Hz), 115.48 (d,  $J$  = 18.5 Hz), 109.58 (d,  $J$  = 3.2 Hz), 107.50 (d,  $J$  = 3.0 Hz), 106.08 (d,  $J$  = 19.6 Hz), 105.44 (d,  $J$  = 5.4 Hz), 53.1.

$^{19}\text{F}$  NMR (565 MHz,  $\text{CDCl}_3$ )  $\delta$  -123.4.

**Methyl (Z)-3-(5-fluoro-1H-indol-3-yl)-2-hydroxyacrylate (B4)**



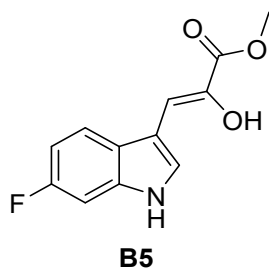
White solid.

$^1\text{H}$  NMR (600 MHz, Chloroform- $d$ )  $\delta$  8.45 (s, 1H), 7.99 (d,  $J$  = 2.7 Hz, 1H), 7.43 (dd,  $J$  = 9.5, 2.5 Hz, 1H), 7.31 (dd,  $J$  = 8.8, 4.3 Hz, 1H), 6.99 (td,  $J$  = 8.9, 2.5 Hz, 1H), 6.84 (d,  $J$  = 1.6 Hz, 1H), 6.28 (d,  $J$  = 1.7 Hz, 1H), 3.93 (s, 3H).

$^{13}\text{C}$  NMR (151 MHz, Chloroform- $d$ )  $\delta$  166.58, 158.53 (d,  $J$  = 236.4 Hz), 136.76, 132.09, 129.04, 127.46 (d,  $J$  = 9.8 Hz), 112.09 (d,  $J$  = 9.8 Hz), 111.21 (d,  $J$  = 26.2 Hz), 110.98 (d,  $J$  = 4.4 Hz), 104.04 (d,  $J$  = 23.9 Hz), 103.79, 53.09.

$^{19}\text{F}$  NMR (565 MHz,  $\text{CDCl}_3$ )  $\delta$  -123.2.

**Methyl (Z)-3-(6-fluoro-1H-indol-3-yl)-2-hydroxyacrylate (B5)**



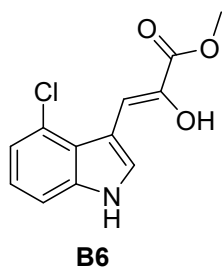
Red solid.

$^1\text{H}$  NMR (400 MHz, Acetonitrile- $d_3$ )  $\delta$  9.64 (s, 1H), 7.88 (d,  $J$  = 2.7 Hz, 1H), 7.72 (dd,  $J$  = 8.7, 5.3 Hz, 1H), 7.19 (dd,  $J$  = 9.9, 2.4 Hz, 1H), 6.94 (ddd,  $J$  = 9.8, 8.7, 2.4 Hz, 1H), 6.88 – 6.82 (m, 2H), 3.85 (s, 3H).

$^{13}\text{C}$  NMR (101 MHz, Acetonitrile- $d_3$ )  $\delta$  166.55, 160.79 (d,  $J$  = 235.4 Hz), 138.31, 136.64 (d,  $J$  = 12.9 Hz), 129.03 (d,  $J$  = 3.1 Hz), 124.29, 120.34 (d,  $J$  = 10.2 Hz), 110.83, 109.18 (d,  $J$  = 24.7 Hz), 104.27, 98.56 (d,  $J$  = 26.2 Hz), 53.12.

$^{19}\text{F}$  NMR (377 MHz,  $\text{CD}_3\text{CN}$ )  $\delta$  -122.7.

**Methyl (Z)-3-(4-chloro-1H-indol-3-yl)-2-hydroxyacrylate (B6)**

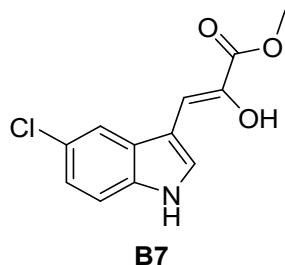


White solid.

$^1\text{H}$  NMR (400 MHz, Chloroform- $d$ )  $\delta$  8.56 (s, 1H), 8.12 (d,  $J$  = 2.8 Hz, 1H), 7.76 (s, 1H), 7.36 – 7.23 (m, 1H), 7.18 – 7.06 (m, 2H), 6.35 (d,  $J$  = 1.7 Hz, 1H).

$^{13}\text{C}$  NMR (101 MHz,  $\text{CDCl}_3$ )  $\delta$  167.0, 137.1, 136.4, 129.2, 126.7, 123.2, 123.1, 122.1, 111.1, 110.3, 105.1, 53.2.

**Methyl (Z)-3-(5-chloro-1H-indol-3-yl)-2-hydroxyacrylate (B7)**

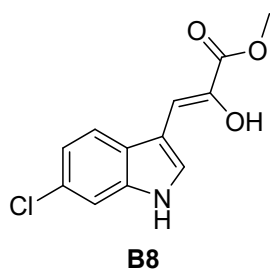


White solid.

$^1\text{H}$  NMR (600 MHz, Chloroform- $d$ )  $\delta$  8.44 (s, 1H), 7.97 (s, 1H), 7.75 (s, 1H), 7.32 (d,  $J$  = 8.6 Hz, 1H), 7.20 (d,  $J$  = 8.5 Hz, 1H), 6.85 (s, 1H), 6.27 (s, 1H), 3.93 (s, 3H).

$^{13}\text{C}$  NMR (151 MHz,  $\text{CDCl}_3$ )  $\delta$  166.5, 137.0, 134.0, 128.5, 128.0, 126.4, 123.2, 118.5, 112.4, 110.6, 103.4, 53.1.

**Methyl (Z)-3-(6-chloro-1H-indol-3-yl)-2-hydroxyacrylate (B8)**

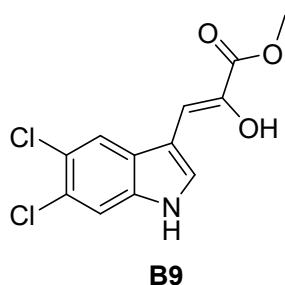


Red solid.

$^1\text{H}$  NMR (600 MHz, Chloroform-*d*)  $\delta$  8.41 (s, 1H), 7.94 (d,  $J$  = 2.7 Hz, 1H), 7.68 (d,  $J$  = 8.4 Hz, 1H), 7.39 (d,  $J$  = 1.8 Hz, 1H), 7.17 (dd,  $J$  = 8.5, 1.8 Hz, 1H), 6.88 (d,  $J$  = 1.5 Hz, 1H), 6.28 (d,  $J$  = 1.6 Hz, 1H), 3.93 (s, 3H).

$^{13}\text{C}$  NMR (151 MHz,  $\text{CDCl}_3$ )  $\delta$  166.5, 137.1, 136.0, 128.7, 127.9, 125.5, 121.3, 119.7, 111.4, 110.9, 103.5, 53.1.

**methyl (Z)-3-(5,6-dichloro-1H-indol-3-yl)-2-hydroxyacrylate (B8)**

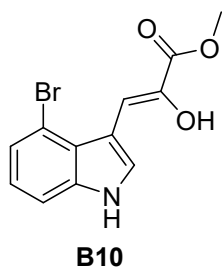


White solid.

$^1\text{H}$  NMR (400 MHz, Chloroform-*d*)  $\delta$  8.43 (s, 1H), 7.96 (d,  $J$  = 2.7 Hz, 1H), 7.85 (s, 1H), 7.51 (s, 1H), 6.79 (d,  $J$  = 1.7 Hz, 1H), 6.31 (d,  $J$  = 1.8 Hz, 1H), 3.93 (s, 3H).

$^{13}\text{C}$  NMR (101 MHz,  $\text{CDCl}_3$ )  $\delta$  166.4, 137.4, 134.4, 128.9, 126.6, 124.8, 120.1, 112.9, 111.7, 110.6, 102.9, 53.2.

**Methyl (Z)-3-(4-bromo-1H-indol-3-yl)-2-hydroxyacrylate (B10)**

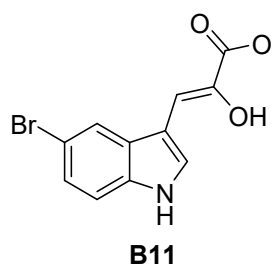


White solid.

$^1\text{H}$  NMR (400 MHz, Chloroform-*d*)  $\delta$  8.57 (s, 1H), 8.15 (d,  $J$  = 2.8 Hz, 1H), 7.93 (s, 1H), 7.40 – 7.30 (m, 2H), 7.05 (t,  $J$  = 7.9 Hz, 1H), 6.35 (d,  $J$  = 1.6 Hz, 1H), 3.93 (s, 3H).

$^{13}\text{C}$  NMR (101 MHz,  $\text{CDCl}_3$ )  $\delta$  167.0, 137.0, 136.1, 129.5, 125.6, 124.3, 123.5, 114.5, 111.5, 110.9, 104.7, 53.2.

**Methyl (Z)-3-(5-bromo-1H-indol-3-yl)-2-hydroxyacrylate (B11)**

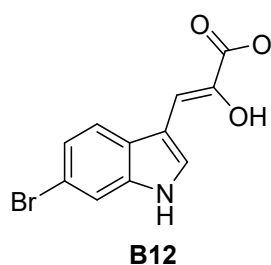


Pink solid.

$^1\text{H}$  NMR (600 MHz, Acetonitrile- $d_3$ )  $\delta$  9.72 (s, 1H), 7.95 (s, 1H), 7.91 (d,  $J$  = 2.5 Hz, 1H), 7.39 (d,  $J$  = 8.6 Hz, 1H), 7.29 (dd,  $J$  = 8.6, 1.5 Hz, 1H), 6.86 (s, 1H), 6.83 (s, 1H), 3.85 (s, 3H).

$^{13}\text{C}$  NMR (151 MHz,  $\text{CD}_3\text{CN}$ )  $\delta$  166.5, 138.4, 135.5, 129.8, 129.4, 125.8, 122.0, 114.3, 113.7, 110.5, 104.1, 53.1.

**Methyl (Z)-3-(6-bromo-1H-indol-3-yl)-2-hydroxyacrylate (B12)**

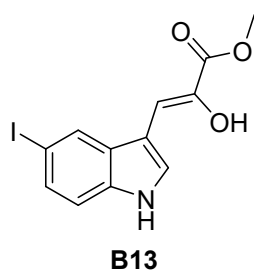


White solid.

$^1\text{H}$  NMR (400 MHz, Chloroform- $d$ )  $\delta$  8.42 (s, 1H), 7.93 (d,  $J$  = 2.6 Hz, 1H), 7.64 (d,  $J$  = 8.5 Hz, 1H), 7.55 (s, 1H), 7.30 (dd,  $J$  = 8.5, 1.7 Hz, 1H), 6.87 (s, 1H), 6.28 (d,  $J$  = 1.8 Hz, 1H), 3.93 (s, 3H).

$^{13}\text{C}$  NMR (101 MHz,  $\text{CDCl}_3$ )  $\delta$  166.5, 137.2, 136.4, 127.8, 125.8, 123.9, 120.1, 116.3, 114.3, 111.0, 103.5, 53.1.

**Methyl (Z)-2-hydroxy-3-(5-iodo-1H-indol-3-yl)acrylate (B13)**

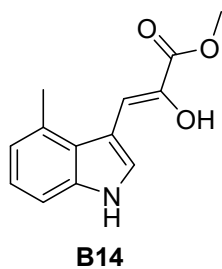


White solid.

$^1\text{H}$  NMR (600 MHz, Chloroform- $d$ )  $\delta$  8.44 (s, 1H), 8.11 (s, 1H), 7.92 (d,  $J$  = 2.6 Hz, 1H), 7.49 (dd,  $J$  = 8.5, 1.6 Hz, 1H), 7.18 (d,  $J$  = 8.4 Hz, 1H), 6.84 (d,  $J$  = 1.6 Hz, 1H), 6.28 (d,  $J$  = 1.7 Hz, 1H), 3.93 (s, 3H).

$^{13}\text{C}$  NMR (151 MHz,  $\text{CDCl}_3$ )  $\delta$  166.5, 137.1, 134.7, 131.2, 129.4, 128.0, 127.8, 113.3, 110.2, 103.3, 84.1, 53.1.

**Methyl (Z)-2-hydroxy-3-(4-methyl-1H-indol-3-yl)acrylate (B14)**

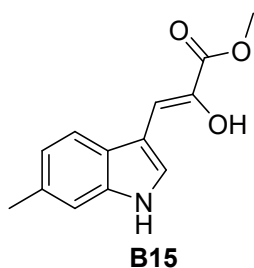


White solid.

$^1\text{H}$  NMR (400 MHz, Chloroform-*d*)  $\delta$  8.43 (s, 1H), 8.09 (s, 1H), 7.31 (s, 1H), 7.26 – 7.24 (m, 1H), 7.12 (t,  $J$  = 7.6 Hz, 1H), 6.94 (d,  $J$  = 7.1 Hz, 1H), 6.29 (s, 1H), 3.93 (s, 3H), 2.80 (s, 3H).

$^{13}\text{C}$  NMR (101 MHz,  $\text{CDCl}_3$ )  $\delta$  166.9, 136.1, 135.9, 131.2, 128.3, 125.1, 122.7, 111.7, 109.5, 106.4, 53.1, 21.3.

**Methyl (Z)-2-hydroxy-3-(6-methyl-1H-indol-3-yl)acrylate (B15)**

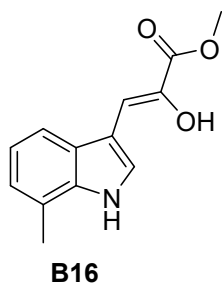


Yellow solid.

$^1\text{H}$  NMR (600 MHz, Chloroform-*d*)  $\delta$  8.29 (s, 1H), 7.90 (d,  $J$  = 2.5 Hz, 1H), 7.67 (d,  $J$  = 8.1 Hz, 1H), 7.19 (s, 1H), 7.04 (dd,  $J$  = 8.0, 1.4 Hz, 1H), 6.94 (d,  $J$  = 1.6 Hz, 1H), 6.22 (d,  $J$  = 1.8 Hz, 1H), 3.93 (s, 3H), 2.48 (s, 3H).

$^{13}\text{C}$  NMR (151 MHz,  $\text{CDCl}_3$ )  $\delta$  166.7, 136.6, 136.1, 132.7, 127.0, 124.8, 122.3, 118.4, 111.3, 110.6, 104.4, 53.0, 21.8.

**Methyl (Z)-2-hydroxy-3-(7-methyl-1H-indol-3-yl)acrylate (B16)**

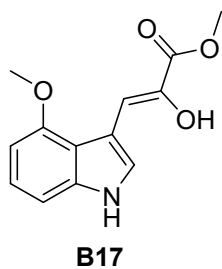


White solid.

$^1\text{H}$  NMR (600 MHz, Chloroform-*d*)  $\delta$  8.35 (s, 1H), 7.96 (d,  $J$  = 2.4 Hz, 1H), 7.65 (d,  $J$  = 7.9 Hz, 1H), 7.14 (t,  $J$  = 7.5 Hz, 1H), 7.06 (d,  $J$  = 7.1 Hz, 1H), 6.97 (s, 1H), 6.27 (s, 1H), 3.93 (s, 3H), 2.51 (s, 3H).

$^{13}\text{C}$  NMR (151 MHz,  $\text{CDCl}_3$ )  $\delta$  166.7, 136.7, 135.2, 127.2, 126.5, 123.4, 120.7, 120.6, 116.4, 111.2, 104.5, 53.0, 16.7.

**Methyl (Z)-2-hydroxy-3-(4-methoxy-1H-indol-3-yl)acrylate (B17)**

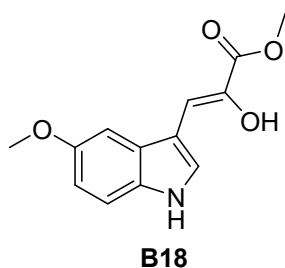


Yellow solid.

$^1\text{H}$  NMR (600 MHz, Chloroform- $d$ )  $\delta$  8.41 (s, 1H), 7.95 (s, 1H), 7.56 (s, 1H), 7.14 (t,  $J$  = 8.0 Hz, 1H), 7.00 (d,  $J$  = 8.1 Hz, 1H), 6.60 (d,  $J$  = 7.8 Hz, 1H), 6.26 (s, 1H), 4.00 (s, 42H), 3.92 (s, 3H).

$^{13}\text{C}$  NMR (151 MHz,  $\text{CDCl}_3$ )  $\delta$  167.0, 155.2, 137.2, 136.1, 126.9, 123.5, 116.3, 111.1, 107.0, 104.7, 100.9, 55.4, 53.0.

**Methyl (Z)-2-hydroxy-3-(5-methoxy-1H-indol-3-yl)acrylate (B18)**

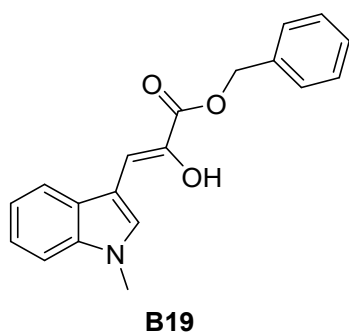


White solid.

$^1\text{H}$  NMR (600 MHz, Acetonitrile- $d_3$ )  $\delta$  9.50 (s, 1H), 7.87 (d,  $J$  = 2.7 Hz, 1H), 7.34 (d,  $J$  = 8.8 Hz, 1H), 7.25 (d,  $J$  = 2.4 Hz, 1H), 6.90 (s, 1H), 6.83 (dd,  $J$  = 8.7, 2.5 Hz, 1H), 6.77 (s, 1H), 3.86 (s, 3H), 3.84 (s, 3H).

$^{13}\text{C}$  NMR (151 MHz,  $\text{CD}_3\text{CN}$ )  $\delta$  166.7, 155.6, 137.6, 131.7, 129.2, 128.2, 113.3, 110.6, 105.0, 100.9, 56.2, 53.0.

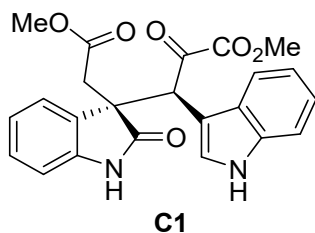
**Benzyl (Z)-2-hydroxy-3-(1-methyl-1H-indol-3-yl)acrylate (B19)**



White solid.

$^1\text{H}$  NMR (400 MHz,  $\text{DMSO}-d_6$ )  $\delta$  9.14 (d,  $J$  = 1.7 Hz, 1H), 7.90 (s, 1H), 7.72 (d,  $J$  = 7.9 Hz, 1H), 7.51 – 7.33 (m, 6H), 7.22 (t,  $J$  = 7.3 Hz, 1H), 7.12 (t,  $J$  = 7.4 Hz, 1H), 6.87 (d,  $J$  = 1.6 Hz, 1H), 5.31 (s, 2H), 3.84 (s, 3H).

$^{13}\text{C}$  NMR (101 MHz,  $\text{DMSO}$ )  $\delta$  164.6, 137.3, 136.4, 136.2, 131.8, 128.6, 128.1, 128.0, 126.8, 122.0, 119.8, 118.3, 110.1, 108.8, 104.9, 66.1, 32.8.



**Methyl (S)-3-(1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (C1)**

Yellow solid; 38.5 mg, 92% yield, >19:1 dr, 96% ee; melting point: 106 – 110 °C;  $[\alpha]_D^{14.6} = -255.8$  ( $c = 0.81$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL OZ-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 8.20 min,  $t_R$  (minor) = 10.68 min.

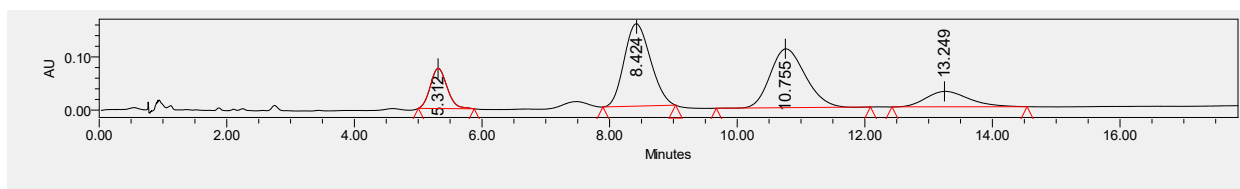
**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.76 (s, 1H), 8.46 (s, 1H), 7.66 (d,  $J = 7.6$  Hz, 1H), 7.35 (d,  $J = 8.0$  Hz, 1H), 7.23 – 7.15 (m, 3H), 6.91 – 6.82 (m, 2H), 6.81 (d,  $J = 2.4$  Hz, 1H), 6.72 (d,  $J = 7.2$  Hz, 1H), 5.50 (s, 1H), 3.54 (s, 3H), 3.33 (s, 3H), 3.15 (dd,  $J = 52.0, 16.0$  Hz, 2H).

**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CDCl}_3$ )  $\delta$  187.8, 180.1, 170.0, 160.9, 142.3, 136.5, 129.3, 128.7, 127.8, 126.9, 125.0, 123.0, 121.7, 120.9, 119.0, 111.9, 110.1, 103.9, 53.2, 51.7, 51.5, 49.9, 40.2.

**IR:** 3365, 2953, 2349, 1620, 1471, 1533, 1470, 1435, 1340, 747  $\text{cm}^{-1}$ .

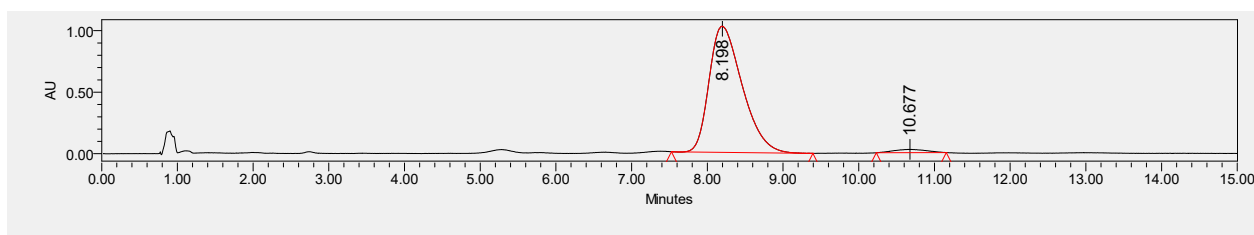
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{23}\text{H}_{20}\text{N}_2\text{O}_6$  419.1249; found 419.1248.

The UPCC chromatograms of racemic product **C1**



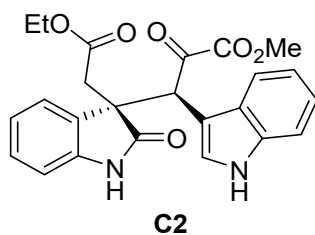
|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 5.312          | 1379872 | 12.09  | 75677  |
| 2 | 8.424          | 4442531 | 38.92  | 155097 |
| 3 | 10.755         | 4234268 | 37.10  | 110819 |
| 4 | 13.249         | 1356487 | 11.89  | 29024  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height  |
|---|----------------|----------|--------|---------|
| 1 | 8.198          | 30985121 | 97.79  | 1022971 |
| 2 | 10.677         | 699846   | 2.21   | 24821   |

**Methyl (S)-3-((S)-3-(2-ethoxy-2-oxoethyl)-2-oxoindolin-3-yl)-3-(1H-indol-3-yl)-2-oxopropanoate (C2)**



Red oil; 38.1 mg, 88% yield, >19:1 dr, 97% ee;  $[\alpha]_D^{14.7} = -192.38$  ( $c = 0.78$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL OZ-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 8.17 min,  $t_R$  (minor) = 10.36 min.

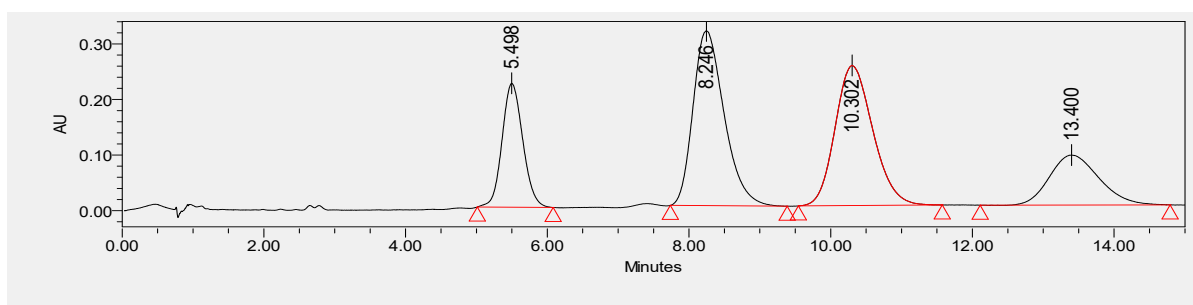
**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.76 (m, 1H), 8.51 (s, 1H), 7.66 (d,  $J = 7.6$  Hz, 1H), 7.35 (d,  $J = 7.9$  Hz, 1H), 7.24 – 7.15 (m, 3H), 6.92 – 6.82 (m, 2H), 6.80 (d,  $J = 4.0$  Hz, 1H), 6.73 (d,  $J = 7.2$  Hz, 1H), 5.49 (s, 1H), 3.84 – 3.72 (m, 2H), 3.54 (s, 3H), 3.13 (dd,  $J = 60.0, 16.0$  Hz, 2H), 0.89 (t,  $J = 8.0$  Hz, 3H).

**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CDCl}_3$ )  $\delta$  187.8, 180.2, 169.4, 160.9, 142.4, 136.5, 129.3, 128.7, 127.8, 126.9, 125.1, 123.0, 121.7, 120.9, 119.0, 111.8, 110.0, 103.9, 60.8, 53.1, 51.6, 50.0, 40.5, 13.8.

**IR:** 3358, 2910, 2850, 2350, 1728, 1620, 1532, 1470, 1433, 747  $\text{cm}^{-1}$ .

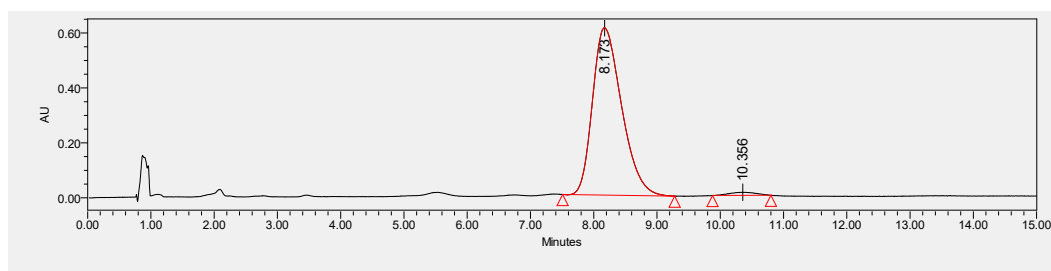
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{24}\text{H}_{22}\text{N}_2\text{O}_6$  433.1405; found 433.1403.

The UPCC chromatograms of racemic product



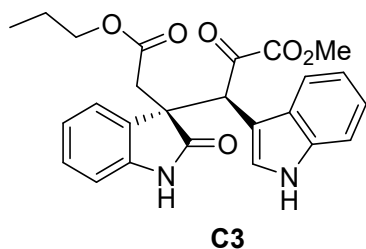
|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 5.498          | 4524009 | 16.28  | 222931 |
| 2 | 8.246          | 9574881 | 34.47  | 314336 |
| 3 | 10.302         | 9309435 | 33.51  | 252258 |
| 4 | 13.400         | 4372666 | 15.74  | 90073  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 8.173          | 19440205 | 98.35  | 609606 |
| 2 | 10.356         | 326417   | 1.65   | 11402  |

**Methyl (S)-3-(1H-indol-3-yl)-2-oxo-3-((S)-2-oxo-3-(2-oxo-2-propoxyethyl)indolin-3-yl)propanoate (C3)**



Yellow solid; 35.8 mg, 80% yield, >19:1 dr, 96% ee; melting point: 139 – 142 °C;  $[\alpha]_D^{14.7} = -192.82$  ( $c = 0.40$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL OZ-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 8.54 min,  $t_R$  (minor) = 10.10 min.

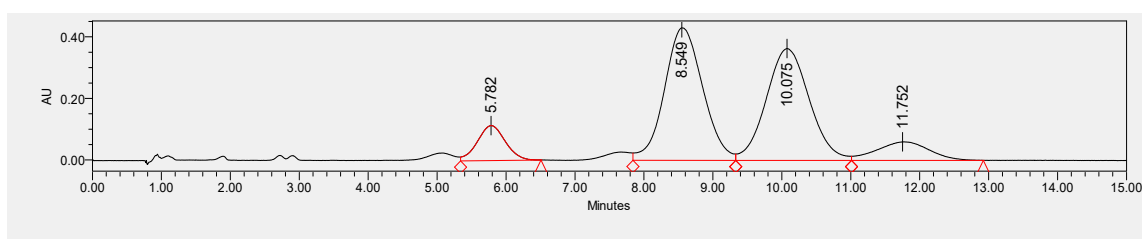
**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.64 (s, 1H), 8.21 (s, 1H), 7.67 (d,  $J = 7.7$  Hz, 1H), 7.39 (s, 1H), 7.25 – 7.16 (m, 3H), 6.92 – 6.80 (m, 3H), 6.72 (d,  $J = 7.4$  Hz, 1H), 5.49 (s, 1H), 3.69 (t,  $J = 6.8$  Hz, 2H), 3.55 (s, 3H), 3.26 – 2.98 (dd,  $J = 58.0, 16.0$  Hz, 2H), 1.34 – 1.28 (m, 2H), 0.70 (t,  $J = 7.4$  Hz, 3H).

**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CDCl}_3$ )  $\delta$  187.8, 180.0, 169.5, 160.9, 142.3, 136.5, 129.3, 128.7, 127.9, 126.8, 125.1, 123.1, 121.7, 121.0, 119.1, 111.8, 109.9, 104.1, 66.4, 53.1, 51.5, 50.0, 40.6, 21.7, 10.3.

**IR**: 3741, 3365, 2960, 2924, 2350, 1728, 1620, 1470, 1433, 748  $\text{cm}^{-1}$ .

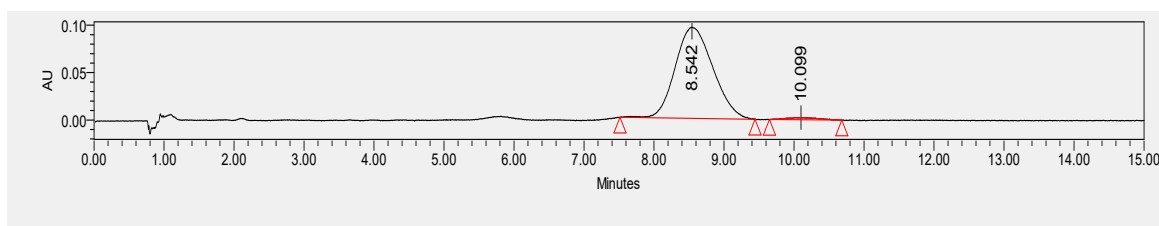
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{25}\text{H}_{24}\text{N}_2\text{O}_6$  447.1562; found 447.1559.

The UPCC chromatograms of racemic product **C3**

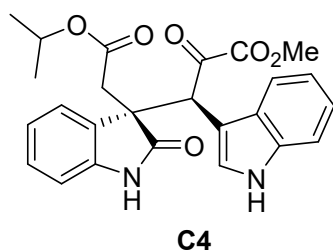


|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 5.782          | 3193277  | 8.22   | 113770 |
| 2 | 8.549          | 16638615 | 42.85  | 430520 |
| 3 | 10.075         | 15794829 | 40.67  | 363374 |
| 4 | 11.752         | 3207365  | 8.26   | 60405  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 8.542          | 3595844 | 98.01  | 96000  |
| 2 | 10.099         | 73027   | 1.99   | 2392   |



**C4**

**Methyl (S)-3-((1H-indol-3-yl)-3-((S)-3-(2-isopropoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (C4)**

Yellow solid; 42.9 mg, 96% yield, >19:1 dr, 93% ee; melting point: 161 – 163 °C;  $[\alpha]_D^{14.8} = -153.25$  ( $c = 0.55$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL OD-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 2.76 min,  $t_R$  (minor) = 3.83 min.

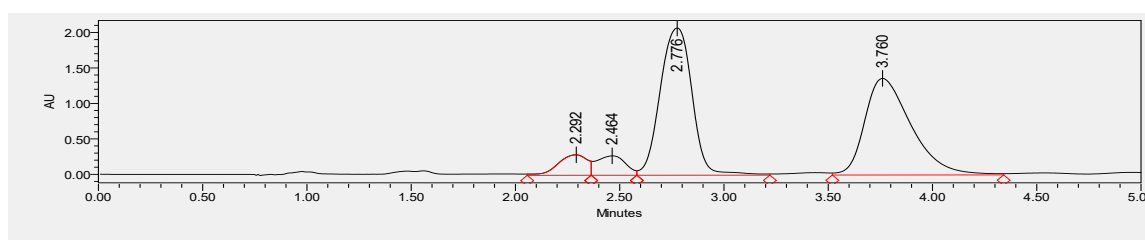
**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.85 (s, 1H), 8.60 (s, 1H), 7.65 (d,  $J = 7.6$  Hz, 1H), 7.35 (d,  $J = 7.6$  Hz, 1H), 7.26 – 7.10 (m, 3H), 6.90 – 6.78 (m, 3H), 6.73 (d,  $J = 7.2$  Hz, 1H), 5.47 (s, 1H), 4.63 (hept,  $J = 6.0$  Hz, 1H), 3.53 (s, 3H), 3.09 (dd,  $J = 68.4, 16.0$  Hz, 2H), 0.86 (dd,  $J = 8.1, 6.4$  Hz, 6H).

**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CDCl}_3$ )  $\delta$  187.8, 180.3, 168.9, 161.0, 142.5, 136.5, 129.2, 128.6, 127.9, 126.9, 125.2, 123.0, 121.6, 120.9, 119.0, 111.9, 110.0, 103.9, 68.4, 53.1, 51.7, 50.1, 40.9, 21.4, 21.2.

**IR**: 3741, 3610, 3365, 2981, 2350, 1728, 1620, 1470, 1262, 749  $\text{cm}^{-1}$ .

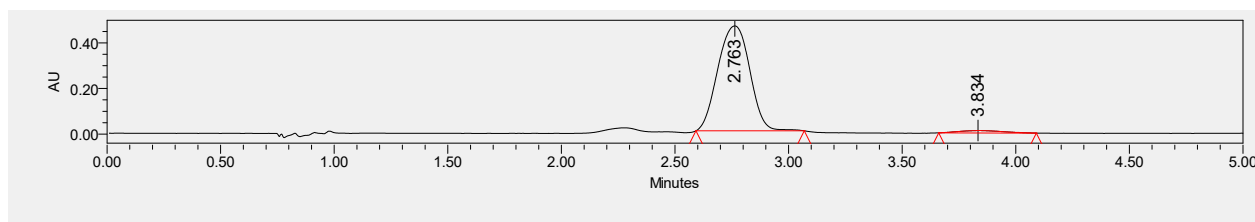
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{25}\text{H}_{24}\text{N}_2\text{O}_6$  447.1562; found 447.1560.

The UPCC chromatograms of racemic product **C4**

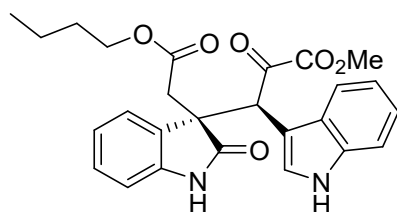


|   | Retention Time | Area     | % Area | Height  |
|---|----------------|----------|--------|---------|
| 1 | 2.292          | 2801181  | 5.76   | 287965  |
| 2 | 2.464          | 2658790  | 5.46   | 272782  |
| 3 | 2.776          | 22065947 | 45.35  | 2075487 |
| 4 | 3.760          | 21133562 | 43.43  | 1362057 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 2.763          | 4513624 | 96.30  | 461116 |
| 2 | 3.834          | 173441  | 3.70   | 11713  |



**C5**

**Methyl (S)-3-((S)-3-(2-butoxy-2-oxoethyl)-2-oxoindolin-3-yl)-3-(1H-indol-3-yl)-2-oxopropanoate (C5)**

Yellow solid; 32.8 mg, 70% yield, >19:1 dr, 95% ee; melting point: 156 – 160 °C;  $[\alpha]_D^{14.7} = -217.98$  ( $c = 0.48$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL OX-3, CO<sub>2</sub>/MeOH = 80/20, flow rate = 1.5 mL/min,  $\lambda$  = 211 nm,  $t_R$  (major) = 8.67 min,  $t_R$  (minor) = 12.56 min.

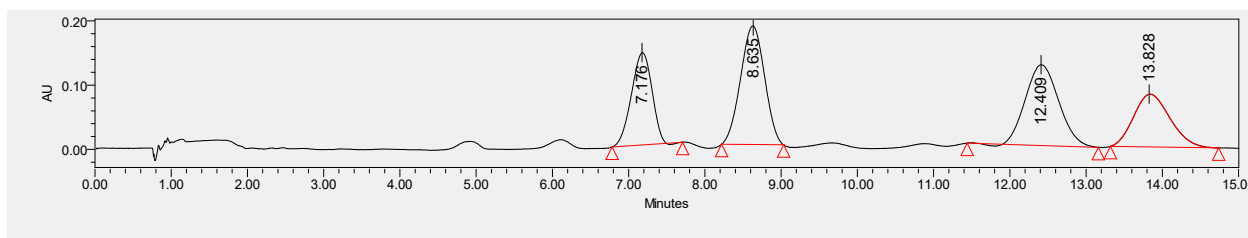
**<sup>1</sup>H NMR** (400 MHz, Chloroform-*d*)  $\delta$  8.70 (s, 1H), 8.31 (s, 1H), 7.67 (d,  $J$  = 7.6 Hz, 1H), 7.36 (d,  $J$  = 7.6 Hz, 1H), 7.25 – 7.16 (m, 3H), 6.922 – 6.79 (m, 3H), 6.73 (d,  $J$  = 7.2 Hz, 1H), 5.49 (s, 1H), 3.74 (t,  $J$  = 6.8 Hz, 2H), 3.55 (s, 3H), 3.12 (dd,  $J$  = 48.0, 16.0 Hz, 2H), 1.33 – 1.19 (m, 2H), 1.15 – 1.04 (m, 2H), 0.76 (t,  $J$  = 7.2 Hz, 3H).

**<sup>13</sup>C NMR** (101 MHz, CDCl<sub>3</sub>)  $\delta$  187.8, 180.0, 169.5, 160.9, 142.3, 136.5, 129.3, 128.7, 127.8, 126.8, 125.0, 123.0, 121.7, 120.9, 119.1, 111.8, 110.0, 104.0, 64.7, 53.1, 51.5, 50.0, 40.6, 30.4, 19.0, 13.7.

**IR**: 3742, 3610, 3360, 2958, 2350, 1729, 1621, 1471, 1262, 749 cm<sup>-1</sup>.

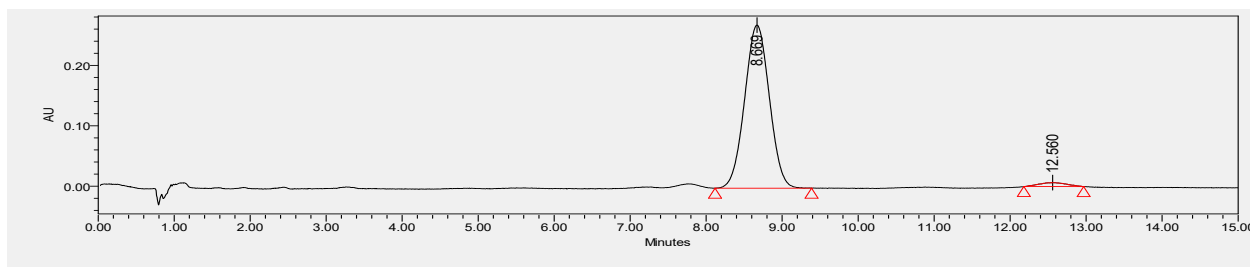
**HRMS** (FTMS+c ESI)  $m/z$ : [M - H]<sup>-</sup> calcd for C<sub>26</sub>H<sub>26</sub>N<sub>2</sub>O<sub>6</sub> 461.1718; found 461.1714.

The UPCC chromatograms of racemic product **C5**

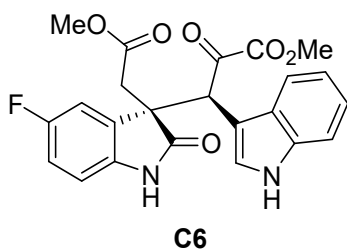


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 7.176          | 2706204 | 20.30  | 143735 |
| 2 | 8.635          | 4029712 | 30.22  | 184608 |
| 3 | 12.409         | 3875042 | 29.06  | 125855 |
| 4 | 13.828         | 2722745 | 20.42  | 82346  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 8.669          | 6095932 | 97.35  | 270073 |
| 2 | 12.560         | 165797  | 2.65   | 6495   |



**Methyl 3-(5-fluoro-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-3-(1H-indol-3-yl)-2-oxopropanoate (C6)**

Yellow solid; 38.0 mg, 87% yield, >19:1 dr, 95% ee; melting point: 111 – 116 °C;  $[\alpha]_D^{14.6} = -242.16$  ( $c = 0.68$  in CH<sub>2</sub>Cl<sub>2</sub>).

**UPCC** DAICEL CHIRALCEL OZ-3, CO<sub>2</sub>/MeOH = 80/20, flow rate = 1.5 mL/min,  $\lambda$  = 211 nm,  $t_R$  (major) = 5.06 min,  $t_R$  (minor) = 5.72 min.

$^1\text{H}$  NMR (400 MHz, Chloroform-*d*)  $\delta$  8.81 (s, 1H), 8.59 (s, 1H), 7.67 (d,  $J$  = 8.0 Hz, 1H), 7.39 – 7.35 (m, 1H), 7.25 – 7.16 (m, 2H), 6.91 (td,  $J$  = 8.0, 2.4 Hz, 1H), 6.84 – 6.77 (m, 2H), 6.47 (dd,  $J$  = 8.0, 4.0 Hz, 1H), 5.50 (s, 1H), 3.56 (s, 3H), 3.36 (s, 3H), 3.13 (dd,  $J$  = 72.0, 16.0 Hz, 2H).

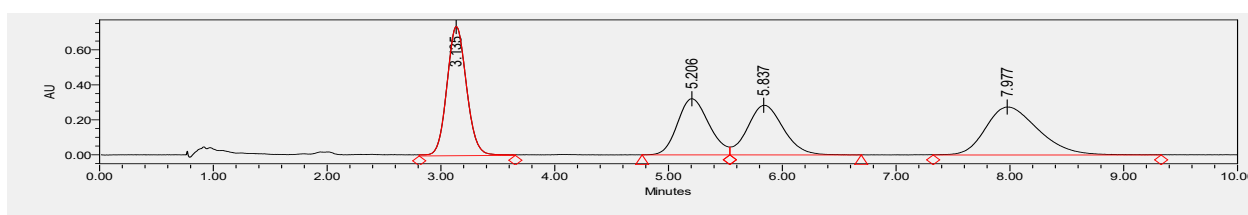
$^{13}\text{C}$  NMR (101 MHz, Chloroform-*d*)  $\delta$  187.71, 180.05, 169.73, 160.82, 158.35 (d,  $J$  = 239.1 Hz), 138.44, 136.60, 131.01 (d,  $J$  = 8.5 Hz), 127.68, 126.77, 123.21, 118.92, 114.94 (d,  $J$  = 23.3 Hz), 112.96 (d,  $J$  = 25.3 Hz), 111.95, 110.49 (d,  $J$  = 8.1 Hz), 103.59, 53.23, 51.90, 51.86, 50.01, 40.22.

$^{19}\text{F}$  NMR (377 MHz,  $\text{CDCl}_3$ )  $\delta$  -121.0.

IR: 3368, 2923, 2350, 1730, 1630, 1486, 1340, 1260, 1198, 747  $\text{cm}^{-1}$ .

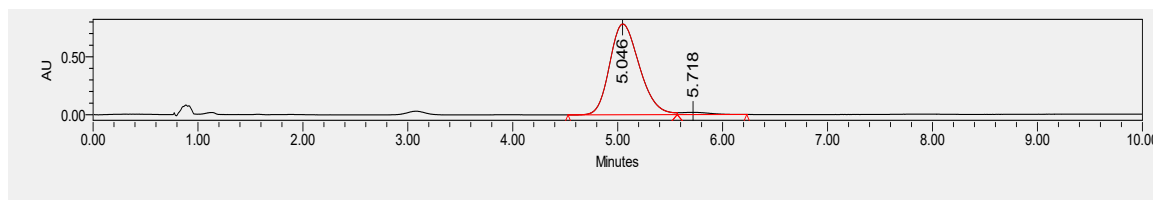
HRMS (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{23}\text{H}_{19}\text{FN}_2\text{O}_6$  437.1154; found 437.1152.

The UPCC chromatograms of racemic product **C6**

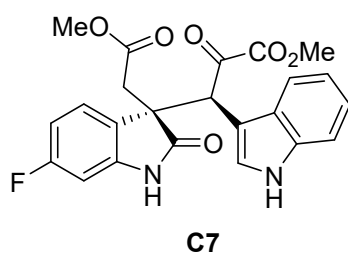


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 3.135          | 8871869 | 29.48  | 738485 |
| 2 | 5.206          | 6187804 | 20.56  | 321236 |
| 3 | 5.837          | 6202977 | 20.61  | 284067 |
| 4 | 7.977          | 8829377 | 29.34  | 274286 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 5.046          | 15404946 | 97.61  | 783354 |
| 2 | 5.718          | 376959   | 2.39   | 19508  |



**Methyl 3-(6-fluoro-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-3-(1H-indol-3-yl)-2-oxopropanoate (C7)**

Yellow oil; 42.0 mg, 96% yield, >19:1 dr, 92% ee;  $[\alpha]_{\text{D}}^{15.0}$  = -229.77 ( $c$  = 0.70 in  $\text{CH}_2\text{Cl}_2$ ).

UPCC DAICEL CHIRALCEL OZ-3,  $\text{CO}_2/\text{MeOH}$  = 80/20, flow rate = 1.5 mL/min,  $\lambda$  = 211 nm,  $t_{\text{R}}$  (major) = 4.70 min,  $t_{\text{R}}$  (minor) = 6.02 min.

$^1\text{H}$  NMR (400 MHz, Chloroform-*d*)  $\delta$  8.83 (d,  $J$  = 2.8 Hz, 1H), 8.77 (s, 1H), 7.65 (d,  $J$  = 7.6 Hz, 1H), 7.36 (d,  $J$  = 7.6 Hz, 1H), 7.20 (m, 2H), 6.80 (d,  $J$  = 2.8 Hz, 1H), 6.66 – 6.59 (m, 2H), 6.53 (ddd,  $J$  = 10.4, 8.4, 2.4 Hz, 1H), 5.47 (s, 1H), 3.55 (s, 3H), 3.35 (s, 3H), 3.15

(dd,  $J = 56.0, 16.0$  Hz, 2H).

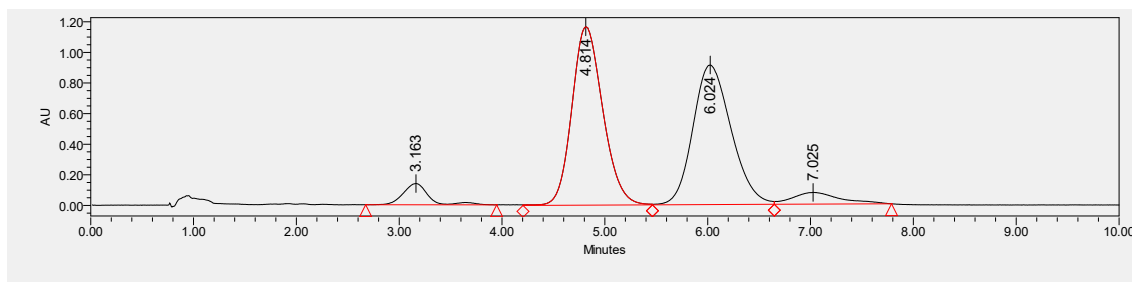
**$^{13}\text{C}$  NMR** (101 MHz, Chloroform- $d$ )  $\delta$  187.89, 180.54, 169.92, 163.17 (d,  $J = 245.2$  Hz), 160.89, 143.87 (d,  $J = 12.0$  Hz), 136.58, 127.66, 126.83, 125.93 (d,  $J = 9.6$  Hz), 124.72 (d,  $J = 2.9$  Hz), 123.13, 120.99, 118.95, 111.93, 107.94 (d,  $J = 22.4$  Hz), 103.67, 98.88 (d,  $J = 27.3$  Hz), 53.21, 51.82, 51.12, 50.01.

**$^{19}\text{F}$  NMR** (377 MHz,  $\text{CDCl}_3$ )  $\delta$  -111.7.

**IR:** 3364, 2954, 2350, 1730, 1628, 1437, 1339, 1264, 1210, 746  $\text{cm}^{-1}$ .

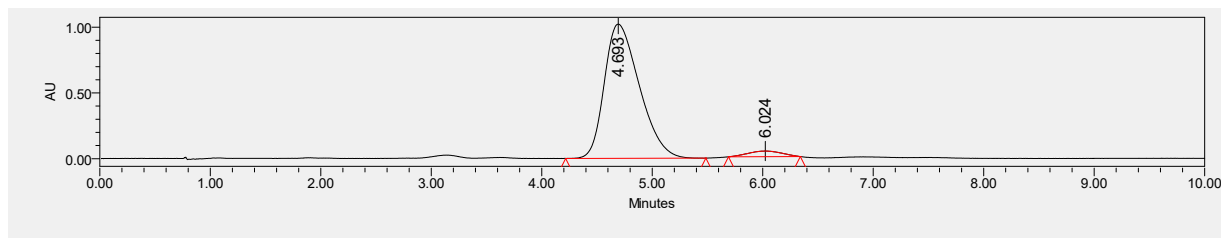
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{23}\text{H}_{19}\text{FN}_2\text{O}_6$  437.1154; found 437.1151.

The UPCC chromatograms of racemic product **C7**

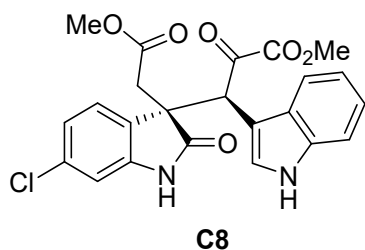


|   | Retention Time | Area     | % Area | Height  |
|---|----------------|----------|--------|---------|
| 1 | 3.163          | 2359905  | 4.48   | 138839  |
| 2 | 4.814          | 24366588 | 46.26  | 1165845 |
| 3 | 6.024          | 23495175 | 44.60  | 912359  |
| 4 | 7.025          | 2452565  | 4.66   | 76330   |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height  |
|---|----------------|----------|--------|---------|
| 1 | 4.693          | 22448099 | 96.09  | 1020136 |
| 2 | 6.024          | 914607   | 3.91   | 43082   |



**Methyl (S)-3-((S)-6-chloro-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-3-(1H-indol-3-yl)-2-oxopropanoate (C8)**

Yellow solid; 43.9 mg, 97% yield, >19:1 dr, 95% ee; melting point: 106 – 114  $^{\circ}\text{C}$ ;  $[\alpha]_{\text{D}}^{14.7} = -200.28$  ( $c = 0.73$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL OX-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_{\text{R}}$  (major) = 6.75 min,  $t_{\text{R}}$  (minor) = 10.89 min.

**$^1\text{H}$  NMR** (400 MHz, Chloroform- $d$ )  $\delta$  8.77 (s, 1H), 8.71 (s, 1H), 7.66 (d,  $J = 7.8$  Hz, 1H), 7.36 (d,  $J = 7.8$  Hz, 1H), 7.24 – 7.17 (m, 2H),

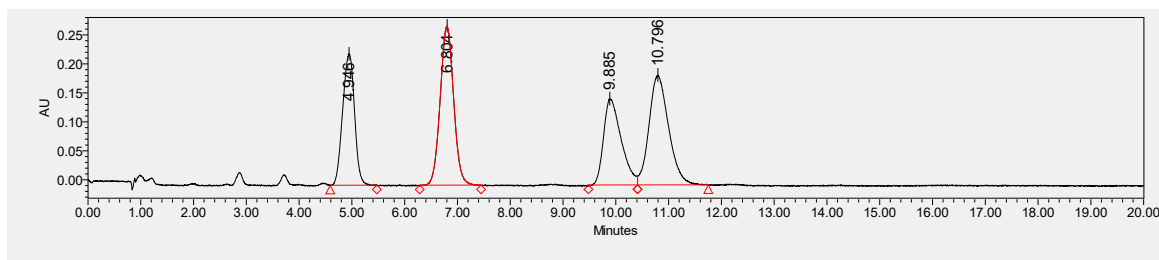
6.90 (d,  $J = 2.0$  Hz, 1H), 6.87 – 6.75 (m, 2H), 6.61 (d,  $J = 8.0$  Hz, 1H), 5.47 (s, 1H), 3.56 (s, 3H), 3.36 (s, 3H), 3.15 (dd,  $J = 56.0, 16.0$  Hz, 2H).

$^{13}\text{C}$  NMR (101 MHz,  $\text{CDCl}_3$ )  $\delta$  187.8, 180.2, 169.8, 160.8, 143.6, 136.6, 134.4, 127.8, 127.7, 126.8, 125.8, 123.2, 121.6, 121.1, 118.9, 111.9, 110.8, 103.7, 53.2, 51.9, 51.2, 50.1, 40.2.  $^{13}\text{C}$  NMR (101 MHz,  $\text{CDCl}_3$ )  $\delta$  187.8, 180.2, 169.8, 160.8, 143.6, 136.6, 134.4, 127.8, 127.7, 126.8, 125.8, 123.2, 121.6, 121.1, 118.9, 111.9, 110.8, 103.7, 53.2, 51.9, 51.2, 50.1, 40.2.

IR: 3365, 2953, 2350, 1730, 1616, 1485, 1337, 1262, 1125, 746  $\text{cm}^{-1}$ .

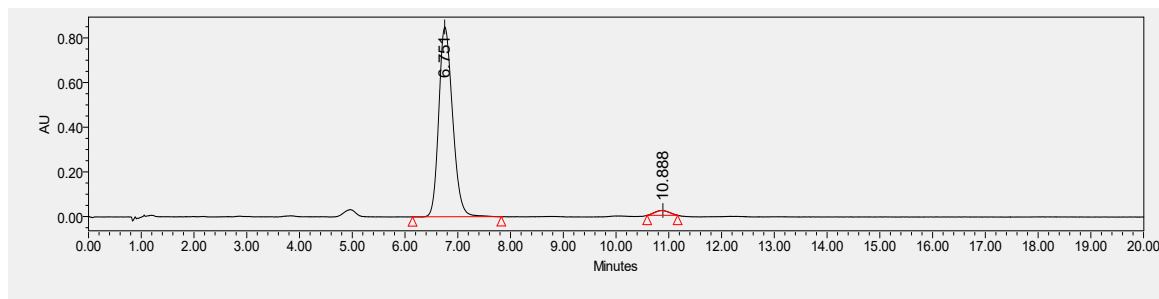
HRMS (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{23}\text{H}_{19}\text{ClN}_2\text{O}_6$  453.0859, 455.0829; found 453.0860, 455.0833.

The UPCC chromatograms of racemic product **C8**

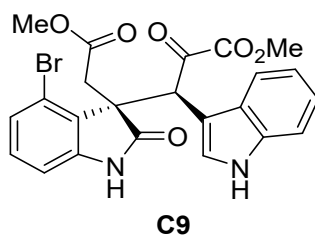


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 4.946          | 3538432 | 21.17  | 226907 |
| 2 | 6.804          | 4857716 | 29.06  | 275052 |
| 3 | 9.885          | 3447818 | 20.63  | 148894 |
| 4 | 10.796         | 4871483 | 29.14  | 189760 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 6.751          | 15438711 | 97.37  | 850182 |
| 2 | 10.888         | 416934   | 2.63   | 21835  |



**Methyl (S)-3-((S)-4-bromo-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-3-(1H-indol-3-yl)-2-oxopropanoate (C9)**

Yellow oil; 34.8 mg, 70% yield, >19:1 dr, 89% ee;  $[\alpha]_{\text{D}}^{14.8} = -217.92$  ( $c = 0.42$  in  $\text{CH}_2\text{Cl}_2$ ).

UPCC DAICEL CHIRALCEL ID-3,  $\text{CO}_2/\text{EtOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_{\text{R}}$  (major) = 3.84 min,  $t_{\text{R}}$  (minor) = 4.29 min.

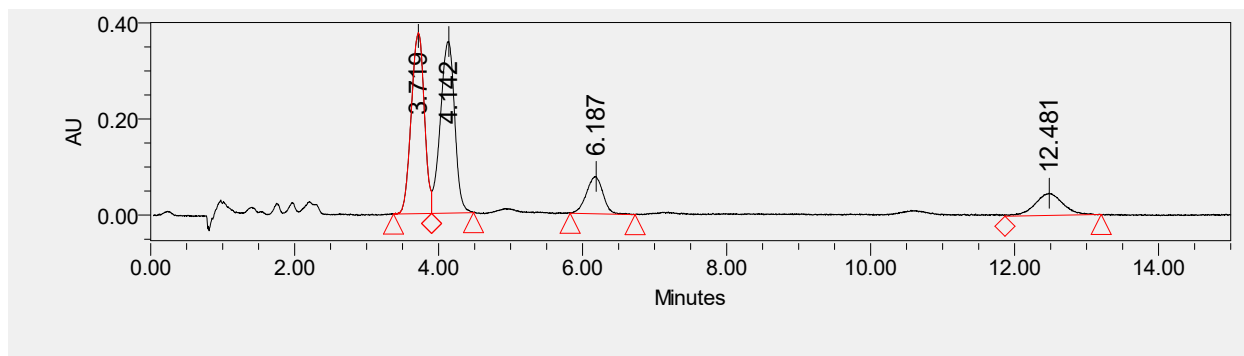
$^1\text{H}$  NMR (400 MHz, Chloroform- $d$ )  $\delta$  8.95 (s, 1H), 8.84 (s, 1H), 7.39 (d,  $J = 7.8$  Hz, 1H), 7.17 (d,  $J = 7.8$  Hz, 1H), 7.07 (d,  $J = 2.8$  Hz, 1H), 7.02 (t,  $J = 8.0$  Hz, 2H), 6.97 – 6.92 (m, 1H), 6.74 (t,  $J = 8.0$  Hz, 1H), 6.35 (d,  $J = 7.6$  Hz, 1H), 5.93 (s, 1H), 3.78 (dd,  $J = 40.0, 16.0$  Hz, 2H), 3.58 (s, 3H), 3.42 (s, 3H).

**<sup>13</sup>C NMR** (101 MHz, CDCl<sub>3</sub>) δ 187.0, 179.0, 170.5, 162.1, 144.0, 135.9, 130.4, 128.3, 127.5, 126.8, 126.6, 122.3, 120.1, 118.3, 118.2, 111.5, 109.1, 103.4, 55.2, 53.2, 52.0, 46.4, 36.8.

**IR:** 3362, 2923, 2349, 1736, 1615, 1450, 1348, 1262, 176, 745 cm<sup>-1</sup>.

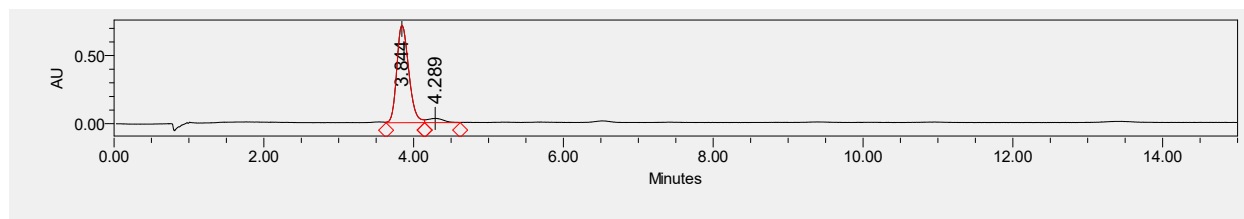
**HRMS** (FTMS+c ESI) m/z: [M - H]<sup>-</sup> calcd for C<sub>23</sub>H<sub>19</sub>BrN<sub>2</sub>O<sub>6</sub> 497.0354, 499.0333; found 497.0355, 499.0334.

The UPCC chromatograms of racemic product **C9**

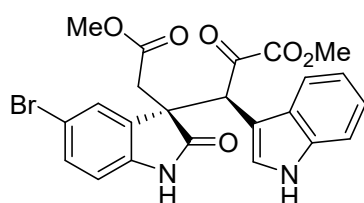


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 3.719          | 4744144 | 38.95  | 376583 |
| 2 | 4.142          | 4943696 | 40.59  | 357036 |
| 3 | 6.187          | 1237132 | 10.16  | 77615  |
| 4 | 12.481         | 1255830 | 10.31  | 46322  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 3.844          | 7948017 | 94.33  | 713874 |
| 2 | 4.289          | 477683  | 5.67   | 31426  |



**C10**

**Methyl (S)-3-((S)-5-bromo-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-3-(1H-indol-3-yl)-2-oxopropanoate (C10)**

Yellow solid; 41.7 mg, 84% yield, >19:1 dr, 96% ee; melting point: 106 – 108 °C; [α]<sub>D</sub><sup>14.7</sup> = -168.06 (c = 0.29 in CH<sub>2</sub>Cl<sub>2</sub>).

**UPCC** DAICEL CHIRALCEL OZ-3, CO<sub>2</sub>/MeOH = 80/20, flow rate = 1.5 mL/min, λ = 211 nm, t<sub>R</sub> (major) = 7.30 min, t<sub>R</sub> (minor) = 8.50 min.

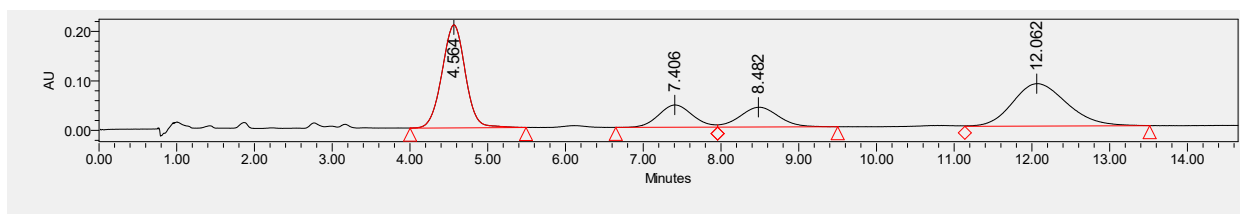
**<sup>1</sup>H NMR** (400 MHz, Chloroform-d) δ 8.64 (s, 1H), 8.27 (s, 1H), 7.67 (d, J = 7.6 Hz, 1H), 7.42 (d, J = 8.0 Hz, 1H), 7.34 (dd, J = 8.0, 2.0 Hz, 1H), 7.29 – 7.20 (m, 2H), 6.85 (d, J = 2.8 Hz, 1H), 6.79 (d, J = 8.0 Hz, 1H), 6.77 (d, J = 2.0 Hz, 1H), 5.48 (s, 1H), 3.57 (s, 3H), 3.39 (s, 3H), 3.10 (dd, J = 94.0, 16.0 Hz, 2H).

**<sup>13</sup>C NMR** (101 MHz, CDCl<sub>3</sub>) δ 187.7, 179.4, 169.6, 160.7, 141.6, 136.6, 131.5, 128.1, 127.6, 126.6, 123.4, 121.2, 119.0, 114.2, 111.9, 111.3, 103.9, 53.2, 51.9, 51.4, 50.2, 40.4.

**IR:** 3365, 2953, 2350, 1730, 1617, 1457, 1338, 1262, 1176, 745 cm<sup>-1</sup>.

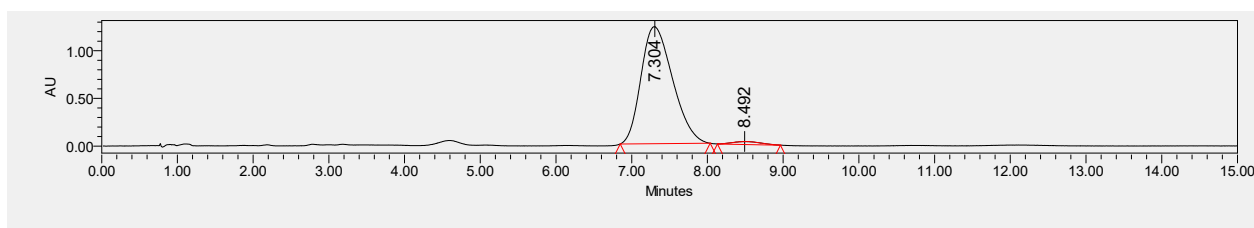
**HRMS** (FTMS+c ESI) m/z: [M - H]<sup>-</sup> calcd for C<sub>23</sub>H<sub>19</sub>BrN<sub>2</sub>O<sub>6</sub> 497.0354, 499.0333; found 497.0354, 499.0334.

The UPCC chromatograms of racemic product **C10**

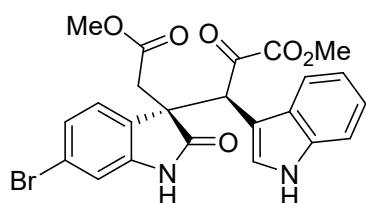


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 4.564          | 4325500 | 38.52  | 208072 |
| 2 | 7.406          | 1368399 | 12.19  | 45039  |
| 3 | 8.482          | 1379286 | 12.28  | 39975  |
| 4 | 12.062         | 4156784 | 37.02  | 85663  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height  |
|---|----------------|----------|--------|---------|
| 1 | 7.304          | 35592730 | 97.80  | 1227024 |
| 2 | 8.492          | 800185   | 2.20   | 31274   |



**C11**

**Methyl (S)-3-((S)-6-bromo-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-3-(1H-indol-3-yl)-2-oxopropanoate (C11)**

Yellow solid; 43.7 mg, 88% yield, >19:1 dr, 96% ee; melting point: 121 – 124 °C; [α]<sub>D</sub><sup>14.6</sup> = -167.98 (c = 0.51 in CH<sub>2</sub>Cl<sub>2</sub>).

**UPCC** DAICEL CHIRALCEL IH-3, CO<sub>2</sub>/MeOH = 85/15, flow rate = 1.5 mL/min, λ = 211 nm, t<sub>R</sub> (major) = 11.35 min, t<sub>R</sub> (minor) = 10.55 min.

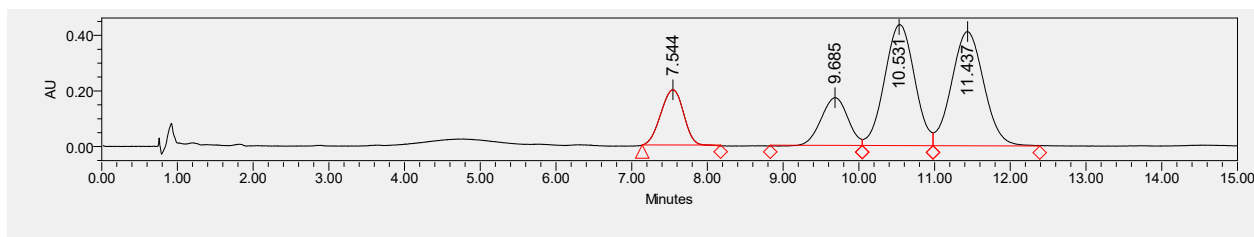
**<sup>1</sup>H NMR** (400 MHz, Chloroform-*d*) δ 8.75 (s, 1H), 8.72 (s, 1H), 7.64 (d, *J* = 8.0 Hz, 1H), 7.37 (d, *J* = 8.0 Hz, 1H), 7.26 – 7.23 (m, 1H), 7.20 (d, *J* = 7.6 Hz, 1H), 7.06 (d, *J* = 1.6 Hz, 1H), 6.99 (dd, *J* = 8.0, 1.6 Hz, 1H), 6.79 (d, *J* = 2.8 Hz, 1H), 6.55 (d, *J* = 8.0 Hz, 1H), 5.46 (s, 1H), 3.56 (s, 3H), 3.36 (s, 3H), 3.15 (dd, *J* = 52.0, 16.0 Hz, 2H).

**<sup>13</sup>C NMR** (101 MHz, CDCl<sub>3</sub>) δ 187.8, 180.1, 169.8, 160.8, 143.8, 136.6, 128.4, 127.6, 126.8, 126.2, 124.6, 123.2, 122.4, 121.1, 118.9, 113.6, 111.9, 103.7, 51.9, 51.2, 50.0, 40.1.

IR: 3366, 2953, 2350, 1730, 1611, 1482, 1337, 1208, 1084, 747  $\text{cm}^{-1}$ .

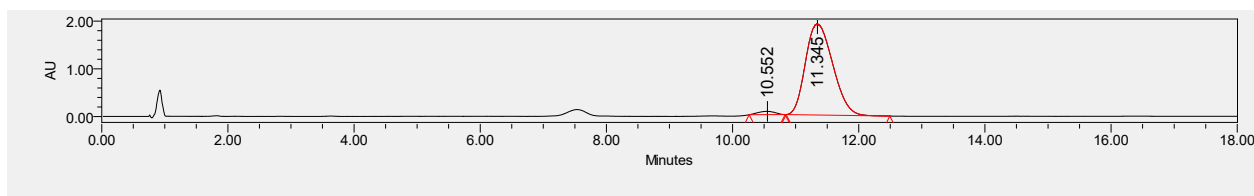
HRMS (FTMS+c ESI)  $m/z$ :  $[M - H]^-$  calcd for  $\text{C}_{23}\text{H}_{19}\text{BrN}_2\text{O}_6$  497.0354, 499.0333; found 497.0352, 499.0332.

The UPCC chromatograms of racemic product **C11**

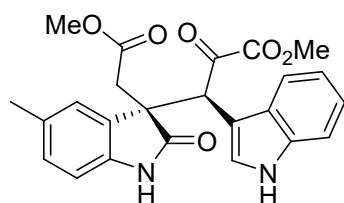


|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 7.544          | 4125502  | 12.71  | 198542 |
| 2 | 9.685          | 4236817  | 13.05  | 171836 |
| 3 | 10.531         | 11940642 | 36.78  | 435195 |
| 4 | 11.437         | 12157885 | 37.45  | 410810 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height  |
|---|----------------|----------|--------|---------|
| 1 | 10.552         | 1320319  | 2.25   | 69699   |
| 2 | 11.345         | 57272650 | 97.75  | 1914363 |



**C12**

**Methyl (S)-3-(1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-5-methyl-2-oxoindolin-3-yl)-2-oxopropanoate (C12)**

Yellow oil; 37.2 mg, 86% yield, >19:1 dr, 98% ee;  $[\alpha]_D^{15.3} = -283.25$  ( $c = 0.4$  in  $\text{CH}_2\text{Cl}_2$ ).

UPCC DAICEL CHIRALCEL OZ-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 9.51 min,  $t_R$  (minor) = 10.94 min.

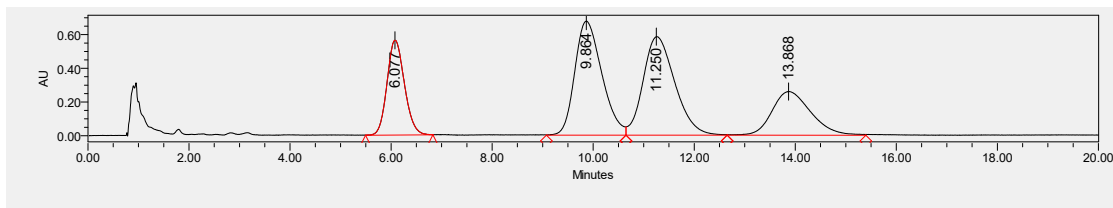
**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.76 (s, 1H), 8.33 (s, 1H), 7.61 (d,  $J = 7.6$  Hz, 1H), 7.34 (d,  $J = 8.0$  Hz, 1H), 7.21 (t,  $J = 7.2$  Hz, 1H), 7.15 (t,  $J = 7.2$  Hz, 1H), 6.99 (d,  $J = 7.6$  Hz, 1H), 6.84 (d,  $J = 2.4$  Hz, 1H), 6.76 (d,  $J = 7.6$  Hz, 1H), 6.54 (s, 1H), 5.47 (s, 1H), 3.55 (s, 3H), 3.36 (s, 3H), 3.14 (dd,  $J = 40.0, 16.0$  Hz, 2H), 2.16 (s, 3H).

**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CDCl}_3$ )  $\delta$  187.9, 180.1, 170.0, 161.0, 139.7, 136.5, 131.0, 129.4, 129.0, 127.8, 126.9, 125.8, 123.0, 120.8, 119.1, 111.8, 109.6, 104.0, 53.1, 51.7, 51.5, 50.0, 40.1, 21.3.

IR: 3367, 2953, 2350, 1730, 1625, 1492, 1339, 1263, 1080, 740  $\text{cm}^{-1}$ .

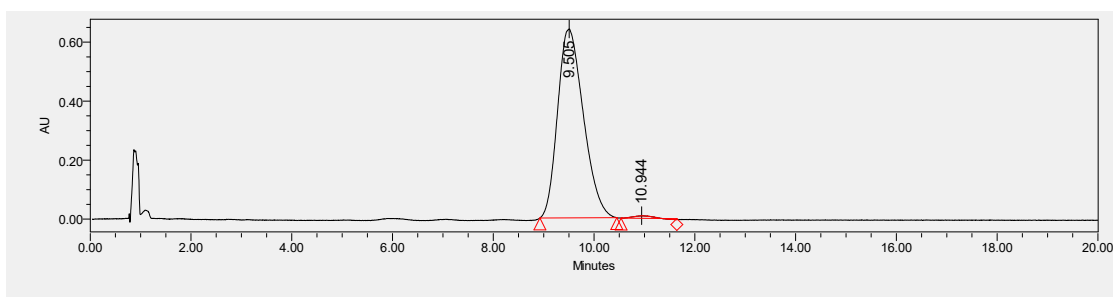
HRMS (FTMS+c ESI)  $m/z$ :  $[M - H]^-$  calcd for  $\text{C}_{24}\text{H}_{22}\text{N}_2\text{O}_6$  433.1405; found 433.1402.

The UPCC chromatograms of racemic product **C12**

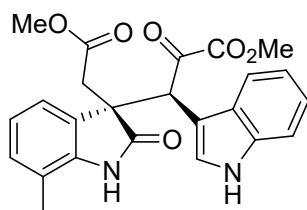


|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 6.077          | 13443692 | 17.24  | 562213 |
| 2 | 9.864          | 25161730 | 32.26  | 677210 |
| 3 | 11.250         | 25504279 | 32.70  | 585137 |
| 4 | 13.868         | 13886016 | 17.80  | 259412 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 9.505          | 22717679 | 98.86  | 640489 |
| 2 | 10.944         | 262670   | 1.14   | 9033   |



**C13**

**Methyl (S)-3-(1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-7-methyl-2-oxoindolin-3-yl)-2-oxopropanoate (C13)**

Yellow oil; 30.7 mg, 71% yield, >19:1 dr, 92% ee;  $[\alpha]_D^{15.5} = -122.50$  ( $c = 0.4$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL OZ-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 6.96 min,  $t_R$  (minor) = 13.62 min.

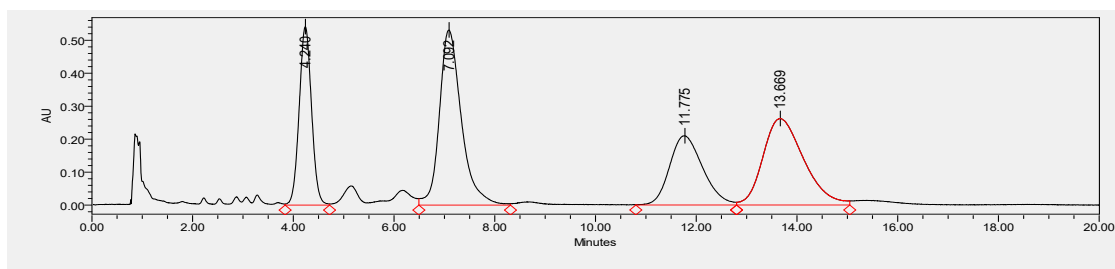
**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.59 (s, 1H), 8.49 (s, 1H), 7.68 (d,  $J = 7.6$  Hz, 1H), 7.37 (d,  $J = 8.0$  Hz, 1H), 7.25 – 7.15 (m, 2H), 7.02 (d,  $J = 7.6$  Hz, 1H), 6.85 (s, 1H), 6.78 (t,  $J = 7.6$  Hz, 1H), 6.56 (d,  $J = 7.4$  Hz, 1H), 3.55 (s, 3H), 3.35 (s, 3H), 3.13 (dd,  $J = 52.0, 16.0$  Hz 2H), 2.24 (s, 3H).

**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CDCl}_3$ )  $\delta$  187.9, 180.3, 169.9, 160.9, 140.9, 136.5, 130.1, 128.9, 127.9, 126.8, 123.1, 122.4, 121.6, 120.9, 119.1, 119.0, 111.7, 104.2, 53.1, 51.8, 51.7, 49.9, 40.2, 16.6.

**IR**: 3366, 2922, 2350, 1730, 1627, 1459, 1437, 1262, 1202, 749  $\text{cm}^{-1}$ .

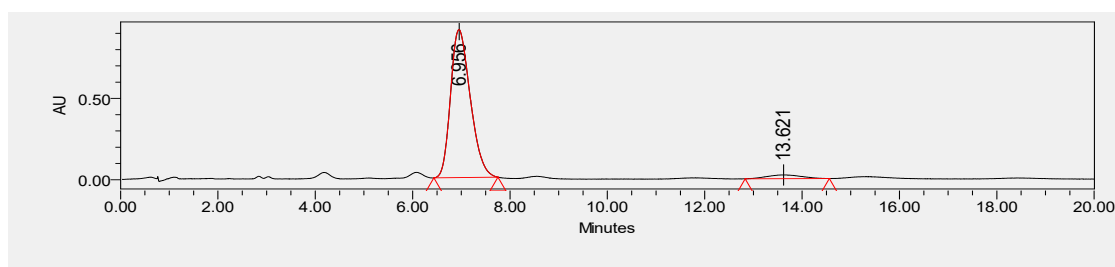
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{24}\text{H}_{22}\text{N}_2\text{O}_6$  433.1405; found 433.1402.

The UPCC chromatograms of racemic product **C13**

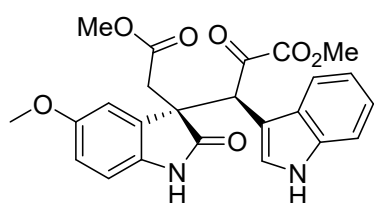


|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 4.240          | 9228088  | 18.60  | 541312 |
| 2 | 7.092          | 16046603 | 32.34  | 531187 |
| 3 | 11.775         | 9607553  | 19.36  | 210006 |
| 4 | 13.669         | 14739329 | 29.70  | 262460 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 6.956          | 25577572 | 95.76  | 910067 |
| 2 | 13.621         | 1131928  | 4.24   | 23270  |



**C14**

**Methyl (S)-3-(1H-indol-3-yl)-3-((S)-5-methoxy-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (C14)**

Yellow oil; 40.9 mg, 91% yield, >19:1 dr, 90% ee;  $[\alpha]_D^{15.6} = -232.07$  ( $c = 0.18$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL IE-3,  $\text{CO}_2/i\text{-PrOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 12.73 min,  $t_R$  (minor) = 11.61 min.

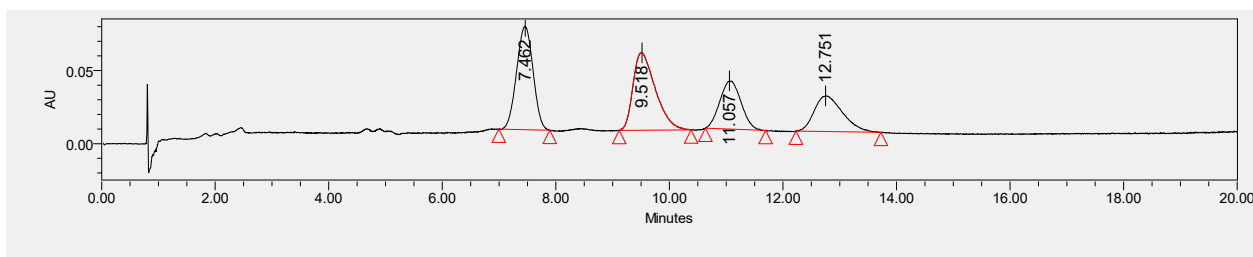
**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.90 (s, 1H), 8.36 (s, 1H), 7.67 (d,  $J = 7.6$  Hz, 1H), 7.35 (d,  $J = 8.0$  Hz, 1H), 7.23 – 7.17 (m, 2H), 6.85 (d,  $J = 2.4$  Hz, 1H), 6.77 – 6.69 (m, 2H), 6.33 (d,  $J = 2.0$  Hz, 1H), 5.50 (s, 1H), 3.60 (s, 3H), 3.55 (s, 3H), 3.36 (s, 3H), 3.11 (dd,  $J = 72.0, 12.0$  Hz, 2H).

**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CDCl}_3$ )  $\delta$  187.9, 180.0, 169.9, 160.9, 154.8, 136.6, 135.9, 130.7, 127.8, 126.9, 123.0, 120.9, 119.0, 112.9, 112.7, 111.9, 110.1, 103.8, 55.7, 53.2, 51.8, 50.1, 40.4.

**IR**: 3364, 2954, 2350, 1731, 1604, 1489, 1340, 1207, 1030, 744  $\text{cm}^{-1}$ .

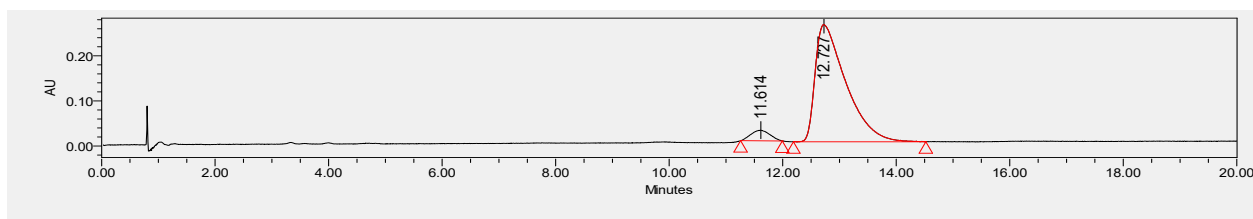
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{24}\text{H}_{22}\text{N}_2\text{O}_7$  449.1354; found 449.1351.

The UPCC chromatograms of racemic product **C14**

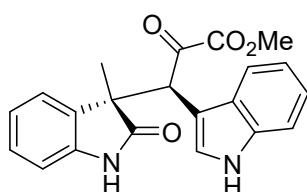


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 7.462          | 1426344 | 31.21  | 70679  |
| 2 | 9.518          | 1447067 | 31.67  | 52971  |
| 3 | 11.057         | 853574  | 18.68  | 32952  |
| 4 | 12.751         | 842593  | 18.44  | 24392  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 11.614         | 540152  | 5.23   | 23669  |
| 2 | 12.727         | 9777976 | 94.77  | 260020 |



**C15**

**Methyl (S)-3-(1H-indol-3-yl)-3-((S)-3-methyl-2-oxindolin-3-yl)-2-oxopropanoate (C15)**

Yellow solid; 28.9 mg, 80% yield, >19:1 dr, 80% ee; melting point: 117 – 120 °C;  $[\alpha]_D^{16.2} = -294.05$  ( $c = 0.37$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL OZ-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 6.09 min,  $t_R$  (minor) = 7.52 min.

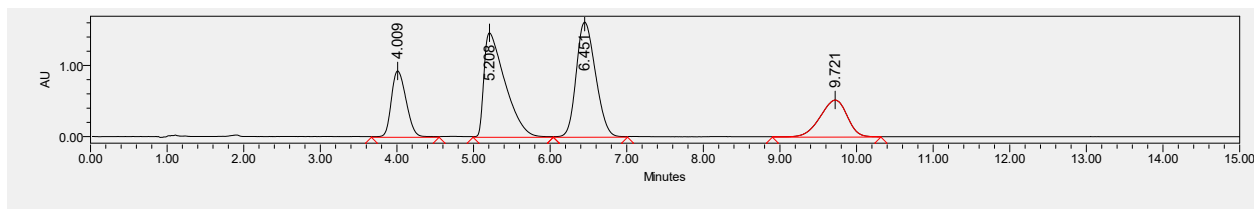
**$^1\text{H}$  NMR** (600 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.65 (s, 1H), 8.47 (s, 1H), 7.76 (d,  $J = 7.6$  Hz, 1H), 7.42 (d,  $J = 8.0$  Hz, 1H), 7.27 (m, 1H), 7.23 – 7.17 (m, 2H), 6.94 (d,  $J = 7.6$  Hz, 1H), 6.83 (t,  $J = 4.0$  Hz, 2H), 6.59 (d,  $J = 7.4$  Hz, 1H), 5.51 (s, 1H), 3.54 (s, 3H), 1.46 (s, 3H).

**$^{13}\text{C}$  NMR** (151 MHz,  $\text{CDCl}_3$ )  $\delta$  189.6, 182.2, 161.2, 141.1, 136.6, 132.5, 128.1, 127.9, 126.2, 124.8, 123.0, 121.7, 121.7, 120.7, 119.6, 111.7, 110.0, 105.2, 53.0, 51.2, 49.7, 24.3.

**IR**: 3370, 2919, 2350, 1727, 1705, 1619, 1470, 1337, 1131, 746  $\text{cm}^{-1}$ .

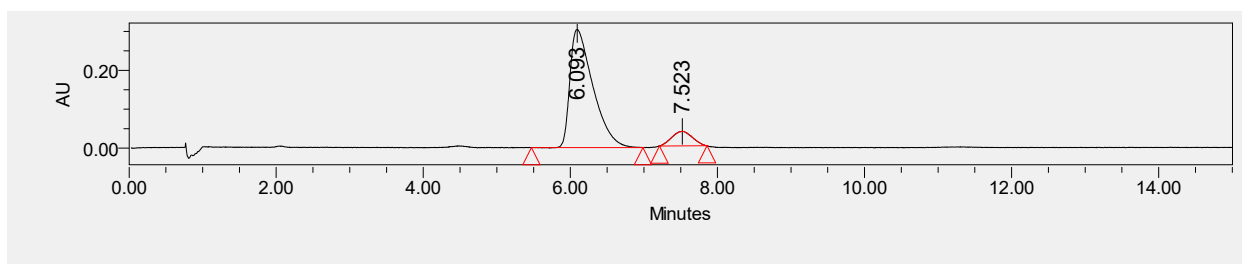
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{21}\text{H}_{18}\text{N}_2\text{O}_4$  361.1194; found 361.1193.

The UPCC chromatograms of racemic product **C15**

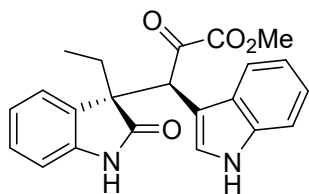


|   | Retention Time | Area     | % Area | Height  |
|---|----------------|----------|--------|---------|
| 1 | 4.009          | 12717305 | 15.44  | 932956  |
| 2 | 5.208          | 28208394 | 34.24  | 1469184 |
| 3 | 6.451          | 28474378 | 34.56  | 1619319 |
| 4 | 9.721          | 12992043 | 15.77  | 521786  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 6.093          | 6591748 | 90.03  | 303359 |
| 2 | 7.523          | 729648  | 9.97   | 37447  |



**C16**

**Methyl (S)-3-((S)-3-ethyl-2-oxoindolin-3-yl)-3-(1H-indol-3-yl)-2-oxopropanoate (C16)**

Yellow solid; 32.0 mg, 85% yield, >19:1 dr, 91% ee; melting point: 90 – 97 °C;  $[\alpha]_D^{16.4} = -265.83$  ( $c = 0.24$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL IH-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 4.15 min,  $t_R$  (minor) = 3.86 min.

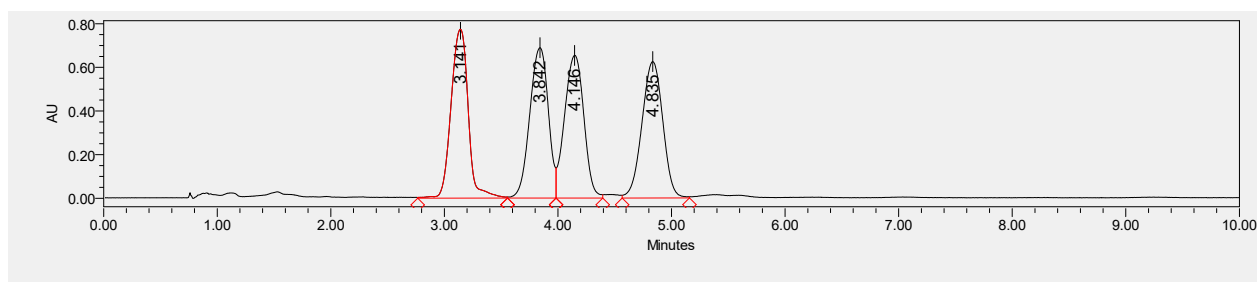
**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.53 (s, 1H), 8.15 (s, 1H), 7.76 (d,  $J = 7.6$  Hz, 1H), 7.42 (d,  $J = 7.6$  Hz, 1H), 7.29 – 7.16 (m, 3H), 6.91 (d,  $J = 7.6$  Hz, 1H), 6.85 (t,  $J = 7.6$  Hz, 1H), 6.81 (d,  $J = 2.4$  Hz, 1H), 6.54 (d,  $J = 7.6$  Hz, 1H), 5.51 (s, 1H), 3.54 (s, 3H), 2.11 (td,  $J = 12.4, 4.4$  Hz, 1H), 1.81 (td,  $J = 12.8, 4.0$  Hz, 1H), 0.67 (t,  $J = 7.2$  Hz, 3H).

**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CDCl}_3$ )  $\delta$  189.3, 181.3, 161.1, 142.0, 136.6, 130.5, 128.1, 127.9, 126.4, 125.1, 123.0, 121.6, 120.8, 119.7, 111.6, 109.6, 105.4, 54.3, 53.0, 51.6, 39.8, 16.8, 14.1.

**IR**: 3370, 2957, 2350, 1727, 1704, 1620, 1468, 1340, 1085, 746  $\text{cm}^{-1}$ .

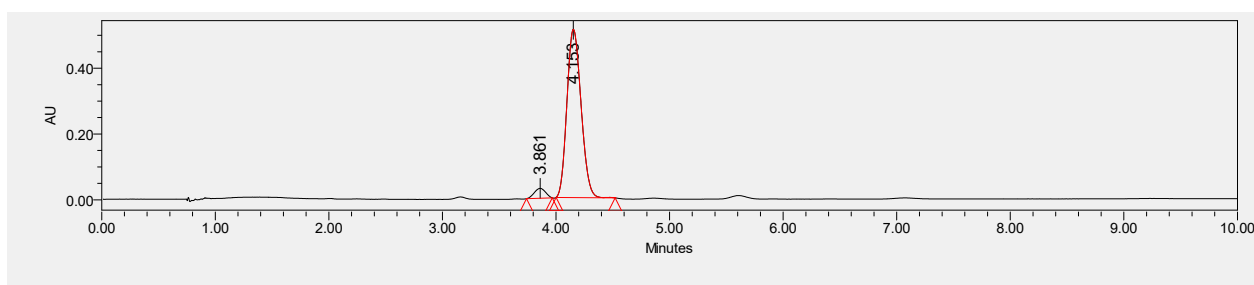
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} + \text{H}]^+$  calcd for  $\text{C}_{22}\text{H}_{20}\text{N}_2\text{O}_4$  377.1496; found 377.1490.

The UPCC chromatograms of racemic product **C16**

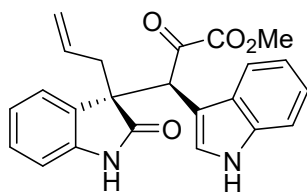


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 3.141          | 8112203 | 25.66  | 773922 |
| 2 | 3.842          | 7806255 | 24.69  | 689015 |
| 3 | 4.146          | 7728095 | 24.45  | 653749 |
| 4 | 4.835          | 7964161 | 25.19  | 625196 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 3.861          | 206471  | 4.46   | 30025  |
| 2 | 4.153          | 4418849 | 95.54  | 511447 |



**C17**

**Methyl (S)-3-((S)-3-allyl-2-oxoindolin-3-yl)-3-(1H-indol-3-yl)-2-oxopropanoate (C17)**

Yellow oil; 31.0 mg, 80% yield, >19:1 dr, 75% ee;  $[\alpha]_D^{16.7} = -277.27$  ( $c = 0.09$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL IG-3,  $\text{CO}_2/\text{MeOH} = 85/15$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 11.90 min,  $t_R$  (minor) = 5.05 min.

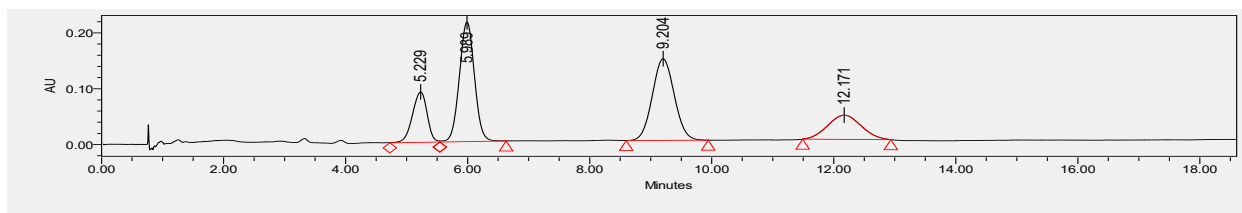
**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.60 (s, 1H), 8.14 (s, 1H), 7.77 (d,  $J = 7.6$  Hz, 1H), 7.43 (d,  $J = 8.0$  Hz, 1H), 7.31 – 7.13 (m, 3H), 6.92 – 6.78 (m, 3H), 6.58 (d,  $J = 7.6$  Hz, 1H), 5.54 (s, 1H), 5.17 – 5.04 (m, 1H), 4.94 – 4.73 (m, 2H), 3.54 (s, 3H), 2.72 (ddd,  $J = 98.4, 13.6, 7.2$  Hz, 2H).

**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CDCl}_3$ )  $\delta$  189.2, 180.6, 161.1, 141.9, 136.6, 131.1, 129.8, 128.1, 128.0, 126.4, 125.3, 123.1, 121.6, 120.9, 119.6, 119.5, 111.7, 109.8, 105.0, 54.2, 53.0, 51.1, 41.9.

**IR**: 3370, 2924, 2406, 1727, 1706, 1620, 1470, 1339, 1261, 747  $\text{cm}^{-1}$ .

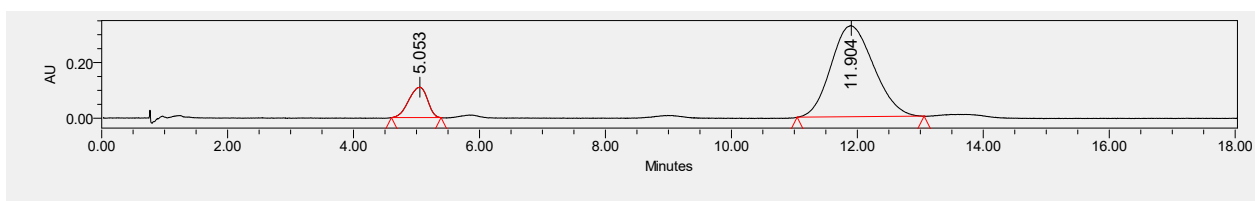
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{23}\text{H}_{20}\text{N}_2\text{O}_4$  387.1350; found 387.1349.

The UPCC chromatograms of racemic product **C17**

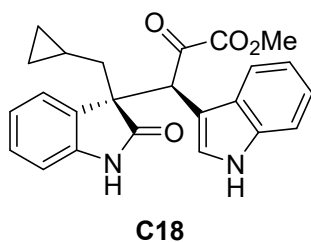


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 5.229          | 1513323 | 14.39  | 90399  |
| 2 | 5.989          | 3679382 | 35.00  | 214244 |
| 3 | 9.204          | 3662722 | 34.84  | 146434 |
| 4 | 12.171         | 1658346 | 15.77  | 43533  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 5.053          | 2237391  | 12.70  | 109333 |
| 2 | 11.904         | 15382448 | 87.30  | 327570 |



**Methyl (S)-3-((S)-3-(cyclopropylmethyl)-2-oxoindolin-3-yl)-3-(1H-indol-3-yl)-2-oxopropanoate (C18)**

Yellow oil; 32.2 mg, 80% yield, >19:1 dr, 87% ee;  $[\alpha]_D^{16.6} = -199.21$  ( $c = 0.38$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL IH-3,  $\text{CO}_2/\text{MeOH} = 85/15$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 9.85 min,  $t_R$  (minor) = 8.22 min.

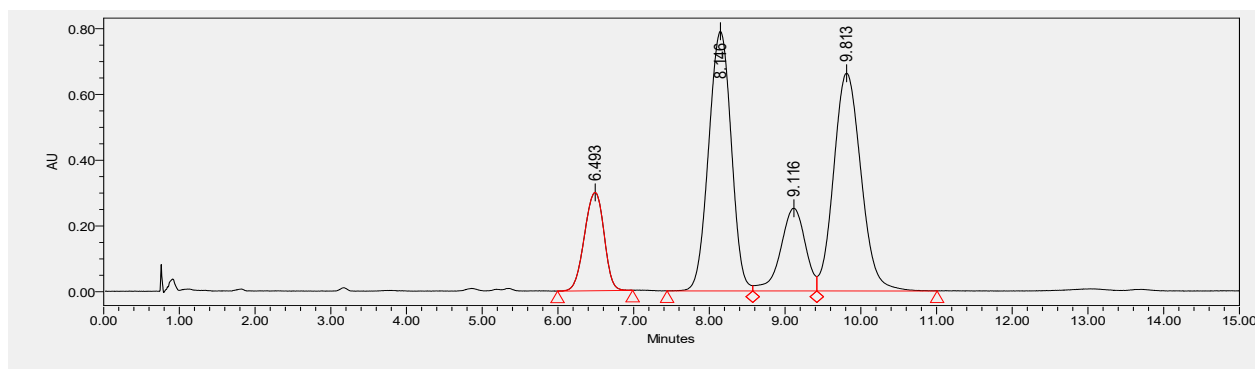
**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.58 (s, 1H), 8.37 (s, 1H), 7.73 (d,  $J = 7.6$  Hz, 1H), 7.42 (d,  $J = 7.8$  Hz, 1H), 7.30 – 7.14 (m, 3H), 6.94 (d,  $J = 7.6$  Hz, 1H), 6.89 – 6.78 (m, 2H), 6.58 (d,  $J = 7.2$  Hz, 1H), 5.48 (s, 1H), 3.54 (s, 3H), 2.12 (dd,  $J = 13.2, 4.8$  Hz, 1H), 1.81 (dd,  $J = 13.2, 8.0$  Hz, 2H), 0.15 – 0.09 (m, 2H), 0.07 – -0.04 (m, 2H), -0.29 (dq,  $J = 9.2, 4.0$  Hz, 1H).

**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CDCl}_3$ )  $\delta$  189.2, 181.8, 161.2, 142.3, 136.6, 130.7, 128.1, 127.9, 126.4, 125.3, 123.0, 121.5, 120.8, 119.6, 111.7, 109.7, 105.2, 54.6, 53.0, 51.4, 42.3, 5.6, 4.1, 3.5.

**IR**: 3371, 2923, 2357, 1728, 1704, 1620, 1469, 1260, 1114, 747  $\text{cm}^{-1}$ .

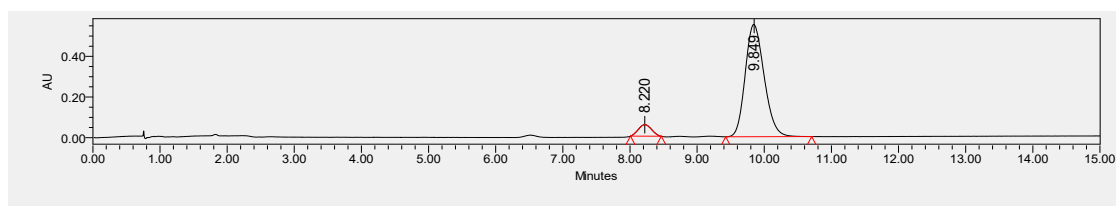
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{24}\text{H}_{22}\text{N}_2\text{O}_4$  401.1057; found 401.1055.

The UPCC chromatograms of racemic product **C18**

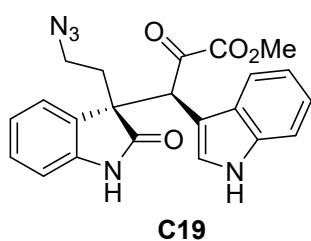


|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 6.493          | 5376493  | 12.20  | 299213 |
| 2 | 8.146          | 16389080 | 37.18  | 789825 |
| 3 | 9.116          | 5691430  | 12.91  | 251463 |
| 4 | 9.813          | 16627202 | 37.72  | 662338 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 8.220          | 772600   | 6.67   | 56506  |
| 2 | 9.849          | 10815945 | 93.33  | 552316 |



**Methyl (S)-3-((S)-3-(2-azidoethyl)-2-oxoindolin-3-yl)-3-(1H-indol-3-yl)-2-oxopropanoate (C19)**

Yellow solid; 39.6 mg, 95% yield, >19:1 dr, 87% ee; melting point: 95 – 104 °C;  $[\alpha]_D^{16.2} = -225.80$  ( $c$

= 0.75 in CH<sub>2</sub>Cl<sub>2</sub>).

**UPCC** DAICEL CHIRALCEL OD-3, CO<sub>2</sub>/MeOH = 90/10, flow rate = 1.5 mL/min,  $\lambda$  = 211 nm,  $t_R$  (major) = 11.91 min,  $t_R$  (minor) = 14.57 min.

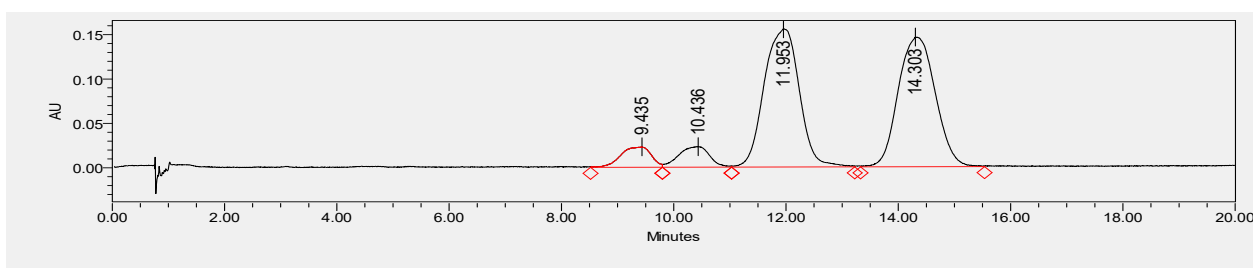
**<sup>1</sup>H NMR** (400 MHz, Chloroform-*d*)  $\delta$  8.61 (s, 1H), 8.46 (s, 1H), 7.73 (d,  $J$  = 7.6 Hz, 1H), 7.42 (d,  $J$  = 7.6 Hz, 1H), 7.32 – 7.19 (m, 3H), 6.96 (d,  $J$  = 7.6 Hz, 1H), 6.88 (t,  $J$  = 7.6 Hz, 1H), 6.82 (d,  $J$  = 2.4 Hz, 1H), 6.58 (d,  $J$  = 7.2 Hz, 1H), 5.51 (s, 1H), 3.55 (s, 3H), 2.90 – 2.74 (m, 2H), 2.47 (dt,  $J$  = 13.6, 8.0 Hz, 1H), 2.14 (ddd,  $J$  = 13.2, 8.0, 4.8 Hz, 2H).

**<sup>13</sup>C NMR** (101 MHz, CDCl<sub>3</sub>)  $\delta$  188.7, 180.5, 161.0, 142.0, 136.6, 128.7, 127.9, 126.5, 125.2, 123.2, 121.9, 121.0, 119.3, 111.8, 110.3, 104.6, 53.1, 52.6, 51.3, 46.9, 36.0.

**IR**: 3369, 2920, 2350, 2100, 1727, 1707, 1620, 1469, 1261, 747 cm<sup>-1</sup>.

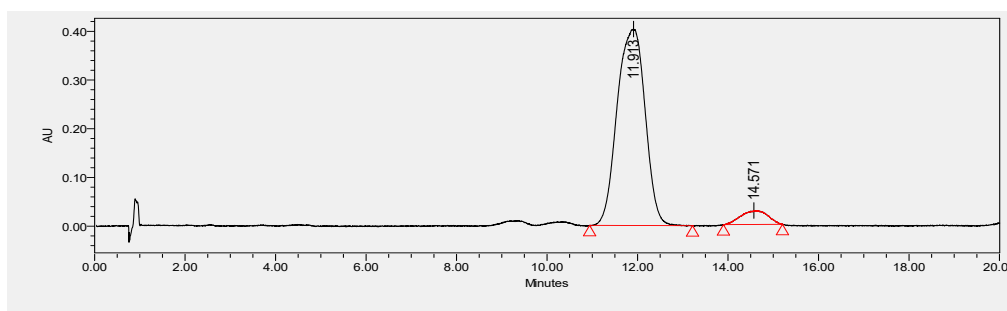
**HRMS** (FTMS+c ESI)  $m/z$ : [M - H]<sup>-</sup> calcd for C<sub>23</sub>H<sub>19</sub>N<sub>5</sub>O<sub>4</sub> 416.1364; found 416.1362.

The UPCC chromatograms of racemic product **C19**

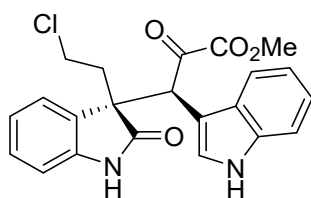


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 9.435          | 863037  | 5.67   | 23114  |
| 2 | 10.436         | 880343  | 5.79   | 23168  |
| 3 | 11.953         | 6765741 | 44.48  | 155738 |
| 4 | 14.303         | 6701973 | 44.06  | 146204 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 11.913         | 17323784 | 93.64  | 403347 |
| 2 | 14.571         | 1176678  | 6.36   | 28820  |



**C20**

**Methyl (S)-3-((S)-3-(2-chloroethyl)-2-oxindolin-3-yl)-3-(1H-indol-3-yl)-2-oxopropanoate (C20)**

Yellow solid; 37.3 mg, 91% yield, >19:1 dr, 82% ee; melting point: 152 – 158 °C;  $[\alpha]_D^{19.8}$  = -249.66 (c

= 0.30 in CH<sub>2</sub>Cl<sub>2</sub>).

**UPCC** DAICEL CHIRALCEL IE-3, CO<sub>2</sub>/MeOH = 85/15, flow rate = 1.5 mL/min,  $\lambda$  = 211 nm,  $t_R$  (major) = 6.00 min,  $t_R$  (minor) = 8.18 min.

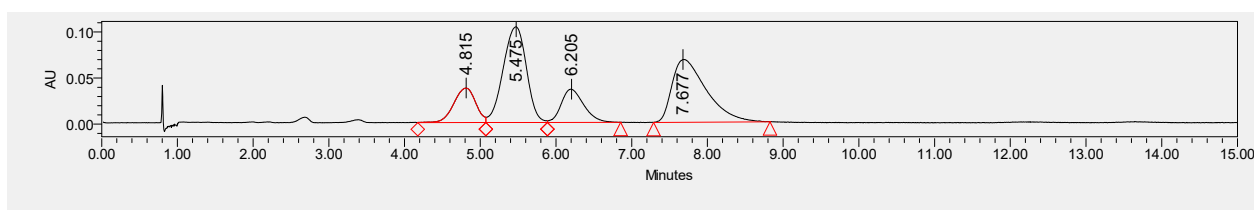
**<sup>1</sup>H NMR** (400 MHz, Chloroform-*d*)  $\delta$  8.67 (s, 1H), 8.62 (s, 1H), 7.72 (d,  $J$  = 7.6 Hz, 1H), 7.42 (d,  $J$  = 7.6 Hz, 1H), 7.30 – 7.20 (m, 3H), 6.94 (d,  $J$  = 7.6 Hz, 1H), 6.88 (t,  $J$  = 7.6 Hz, 1H), 6.82 (d,  $J$  = 2.4 Hz, 1H), 6.59 (d,  $J$  = 7.2 Hz, 1H), 5.52 (s, 1H), 3.54 (s, 3H), 3.14 (td,  $J$  = 10.8, 6.0 Hz, 1H), 2.82 (td,  $J$  = 10.8, 4.8 Hz, 1H), 2.66 (ddd,  $J$  = 13.2, 10.4, 6.0 Hz, 1H), 2.38 (ddd,  $J$  = 13.2, 10.8, 4.4 Hz, 1H).

**<sup>13</sup>C NMR** (101 MHz, CDCl<sub>3</sub>)  $\delta$  188.7, 180.4, 161.0, 142.0, 136.6, 128.7, 128.6, 127.8, 126.6, 125.3, 123.2, 122.0, 121.0, 119.3, 111.8, 110.3, 104.4, 53.3, 53.1, 51.2, 39.7, 39.3.

**IR:** 3369, 2919, 2357, 1727, 1708, 1620, 1470, 1261, 1084, 748 cm<sup>-1</sup>.

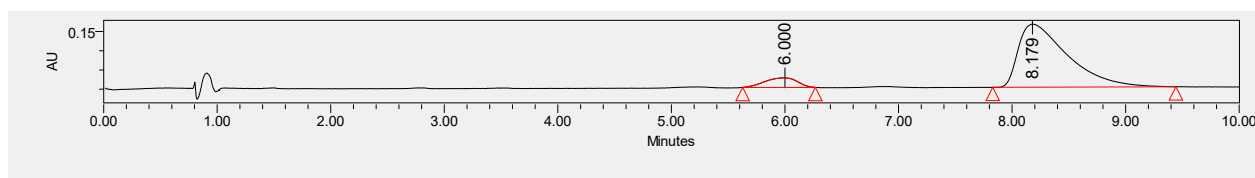
**HRMS** (FTMS+c ESI)  $m/z$ : [M - H]<sup>-</sup> calcd for C<sub>22</sub>H<sub>19</sub>ClN<sub>2</sub>O<sub>4</sub> 409.0961, 411.0931; found 409.0958, 411.0926.

The UPCC chromatograms of racemic product **C20**

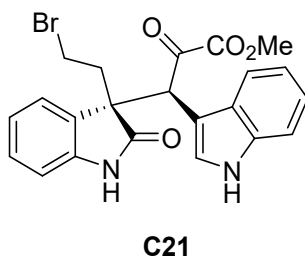


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 4.815          | 750157  | 12.80  | 37521  |
| 2 | 5.475          | 2213905 | 37.79  | 103914 |
| 3 | 6.205          | 746189  | 12.74  | 36131  |
| 4 | 7.677          | 2148637 | 36.67  | 68127  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 6.000          | 496395  | 9.12   | 24890  |
| 2 | 8.179          | 4947310 | 90.88  | 163967 |



**Methyl (S)-3-((S)-3-(2-bromoethyl)-2-oxoindolin-3-yl)-3-(1H-indol-3-yl)-2-oxopropanoate (C21)**

Yellow solid; 40.9 mg, 90% yield, >19:1 dr, 93% ee; melting point: 164 – 166 °C; [ $\alpha$ ]<sub>D</sub><sup>19.8</sup> = -190.73 ( $c$  = 0.25 in CH<sub>2</sub>Cl<sub>2</sub>).

**UPCC** DAICEL CHIRALCEL IE-3, CO<sub>2</sub>/MeOH = 85/15, flow rate = 1.5 mL/min,  $\lambda$  = 211 nm,  $t_R$  (major) = 10.37 min,  $t_R$  (minor) = 7.73 min.

**<sup>1</sup>H NMR** (400 MHz, Chloroform-*d*)  $\delta$  8.67 (s, 1H), 8.64 (s, 1H), 7.71 (d,  $J$  = 7.6 Hz, 1H), 7.42 (d,  $J$  = 7.6 Hz, 1H), 7.32 – 7.19 (m, 3H), 6.94 (d,  $J$  = 7.6 Hz, 1H), 6.88 (t,  $J$  = 7.6 Hz, 1H), 6.83 (d,  $J$  = 2.4 Hz, 1H), 6.60 (d,  $J$  = 7.6 Hz, 1H),

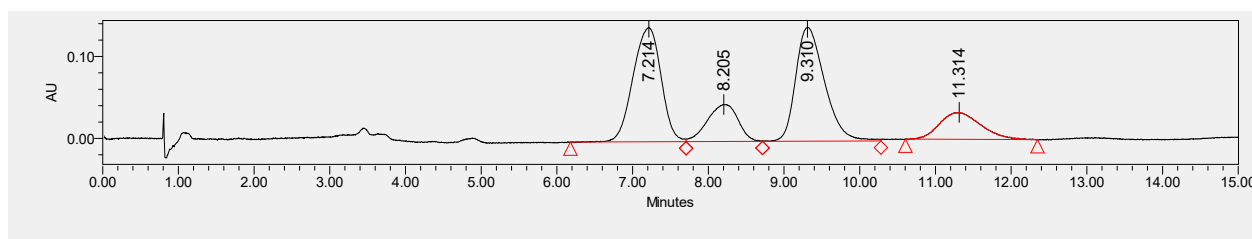
5.51 (s, 1H), 3.54 (s, 2H), 3.00 (ddd,  $J = 12.0, 9.6, 5.2$  Hz, 1H), 2.74 (td,  $J = 12.4, 12.0, 5.2$  Hz, 1H), 2.61 (td,  $J = 11.2, 10.4, 4.4$  Hz, 1H), 2.46 (td,  $J = 12.4, 4.4$  Hz, 1H).

$^{13}\text{C}$  NMR (101 MHz,  $\text{CDCl}_3$ )  $\delta$  188.7, 180.2, 161.0, 142.0, 136.6, 128.7, 128.6, 127.8, 126.6, 125.3, 123.2, 122.1, 121.0, 119.3, 111.8, 110.3, 104.4, 54.3, 53.1, 51.2, 40.1, 26.5.

IR: 3369, 2955, 2349, 1727, 1707, 1620, 1469, 1470, 1261, 746  $\text{cm}^{-1}$ .

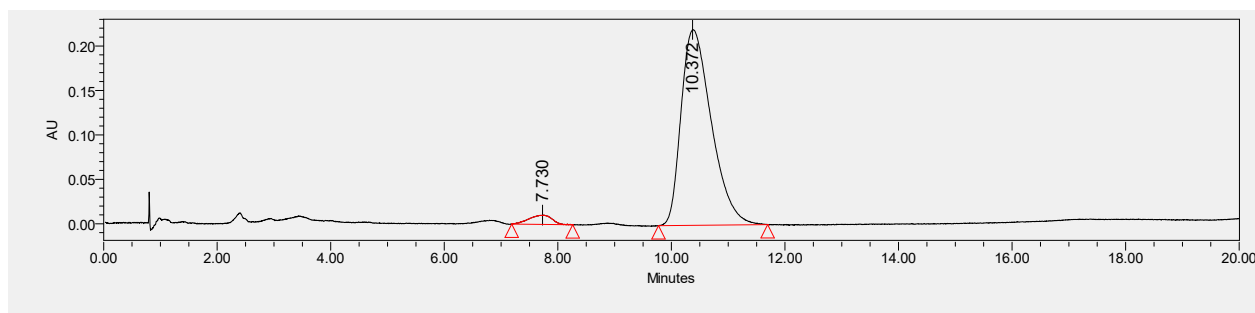
HRMS (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{22}\text{H}_{19}\text{BrN}_2\text{O}_4$  453.0455, 455.0435; found 453.0453, 455.0429.

The UPCC chromatograms of racemic product **C21**

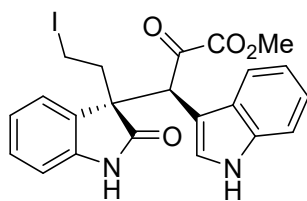


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 7.214          | 3580596 | 36.66  | 139929 |
| 2 | 8.205          | 1266422 | 12.97  | 45192  |
| 3 | 9.310          | 3682189 | 37.70  | 139077 |
| 4 | 11.314         | 1238577 | 12.68  | 32901  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 7.730          | 292043  | 3.46   | 10517  |
| 2 | 10.372         | 8157135 | 96.54  | 220500 |



**C22**

**Methyl (S)-3-(1H-indol-3-yl)-3-((S)-3-(2-iodoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (C22)**

Yellow solid; 46.7 mg, 93% yield, >19:1 dr, 92% ee; melting point: 143 – 153 °C;  $[\alpha]_{\text{D}}^{19.8} = -169.91$  ( $c = 0.68$  in  $\text{CH}_2\text{Cl}_2$ ).

UPCC DAICEL CHIRALCEL IE-3,  $\text{CO}_2/\text{MeOH} = 85/15$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_{\text{R}}$  (major) = 9.77 min,  $t_{\text{R}}$  (minor) = 7.90 min.

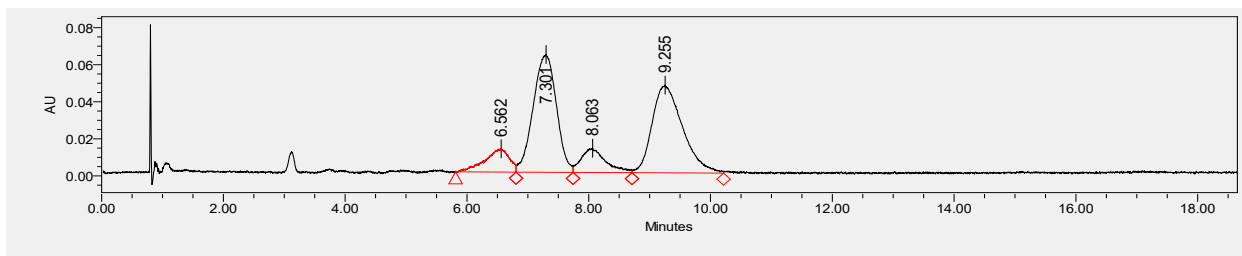
$^1\text{H}$  NMR (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.63 (s, 1H), 8.59 (s, 1H), 7.70 (d,  $J = 7.6$  Hz, 1H), 7.42 (d,  $J = 7.6$  Hz, 1H), 7.35 – 7.17 (m, 3H), 6.94 (d,  $J = 7.6$  Hz, 1H), 6.87 (t,  $J = 7.6$  Hz, 1H), 6.82 (d,  $J = 2.4$  Hz, 1H), 6.58 (d,  $J = 7.2$  Hz, 1H), 5.50 (s, 1H), 3.54 (s, 3H), 2.88 – 2.70 (m, 2H), 2.48 (td,  $J = 13.2, 12.4, 4.4$  Hz, 1H), 2.36 (ddd,  $J = 14.4, 9.2, 4.4$  Hz, 1H).

$^{13}\text{C}$  NMR (101 MHz,  $\text{CDCl}_3$ )  $\delta$  188.7, 180.0, 160.9, 142.0, 136.6, 128.6, 128.5, 127.7, 126.5, 125.3, 123.2, 122.1, 121.1, 119.3, 111.8, 110.3, 104.4, 55.8, 53.1, 51.0, 41.6, -2.9.

IR: 3371, 2952, 2349, 1727, 1707, 1620, 1469, 1261, 1016, 747  $\text{cm}^{-1}$ .

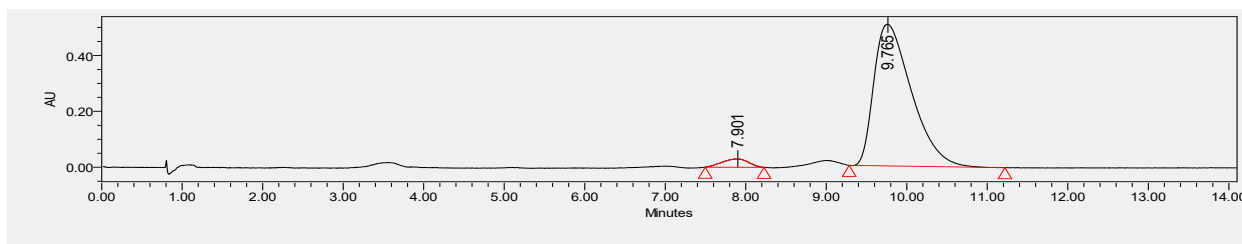
HRMS (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{22}\text{H}_{19}\text{N}_2\text{O}_4$  501.0317; found 501.0311.

The UPCC chromatograms of racemic product **C22**

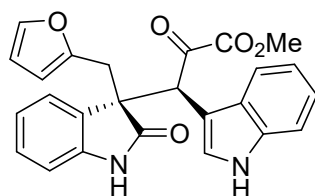


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 6.562          | 346696  | 8.63   | 12592  |
| 2 | 7.301          | 1676971 | 41.74  | 63507  |
| 3 | 8.063          | 359310  | 8.94   | 13111  |
| 4 | 9.255          | 1634867 | 40.69  | 47349  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 7.901          | 677231   | 3.95   | 29982  |
| 2 | 9.765          | 16485416 | 96.05  | 506892 |



**C23**

**Methyl (S)-3-((S)-3-(furan-2-ylmethyl)-2-oxoindolin-3-yl)-3-(1H-indol-3-yl)-2-oxopropanoate (C23)**

Yellow solid; 35.1 mg, 82% yield, >19:1 dr, 90% ee; melting point: 110 – 115  $^{\circ}\text{C}$ ;  $[\alpha]_{\text{D}}^{19.9} = -168.60$  ( $c = 0.48$  in  $\text{CH}_2\text{Cl}_2$ ).

UPCC DAICEL CHIRALCEL IG-3,  $\text{CO}_2/\text{MeOH} = 85/15$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_{\text{R}}$  (major) = 15.10 min,  $t_{\text{R}}$  (minor) = 8.18 min.

$^1\text{H}$  NMR (400 MHz,  $\text{Chloroform-d}$ )  $\delta$  8.55 (s, 1H), 7.81 – 7.72 (m, 2H), 7.45 (d,  $J = 8.0$  Hz, 1H), 7.31 – 7.20 (m, 2H), 7.12 (t,  $J = 7.6$  Hz, 1H), 6.99 (d,  $J = 1.2$  Hz, 1H), 6.89 (d,  $J = 2.4$  Hz, 1H), 6.84 (t,  $J = 7.6$  Hz, 1H), 6.75 (d,  $J = 7.6$  Hz, 1H), 6.66 (d,  $J = 7.2$  Hz, 1H), 5.96 (dd,  $J = 3.2, 1.8$  Hz, 1H), 5.61 (s, 1H), 5.52 (d,  $J = 3.2$  Hz, 1H), 3.56 (s, 3H), 3.35 (dd,  $J = 128.4, 14.4$  Hz, 2H).

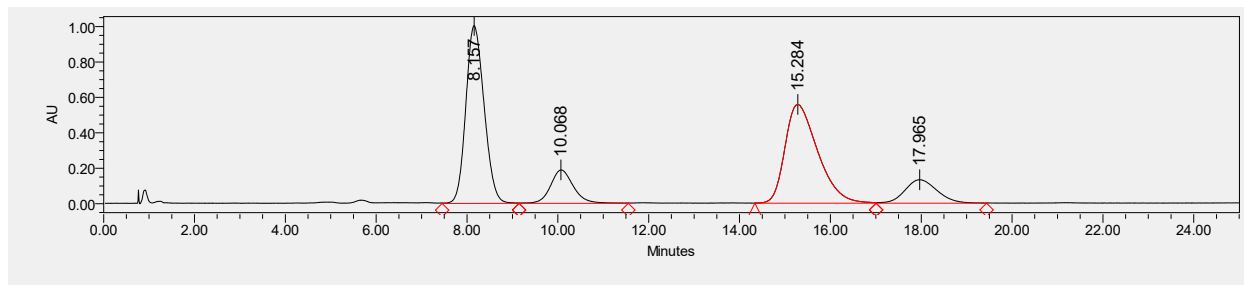
$^{13}\text{C}$  NMR (101 MHz,  $\text{CDCl}_3$ )  $\delta$  188.8, 180.2, 161.1, 149.7, 141.8, 141.5, 136.6, 129.5, 128.2, 127.9, 126.5, 125.6, 123.2, 121.5, 121.0,

119.5, 111.7, 110.0, 109.6, 107.7, 104.9, 54.2, 53.0, 50.7, 36.0.

IR: 3367, 2918, 2350, 1728, 1708, 1620, 1470, 1261, 1014, 747  $\text{cm}^{-1}$ .

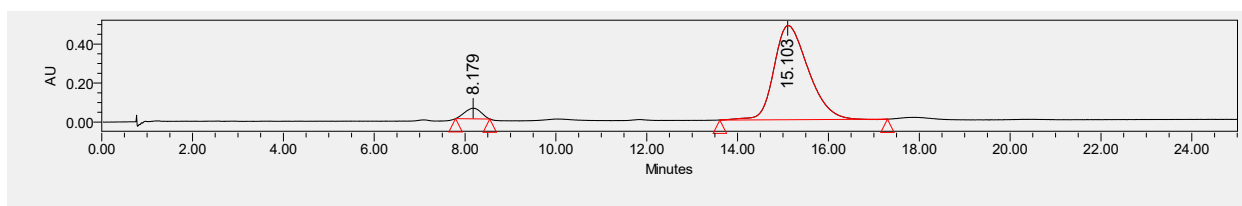
HRMS (FTMS+c ESI)  $m/z$ :  $[M - H]^-$  calcd for  $\text{C}_{25}\text{H}_{20}\text{N}_2\text{O}_5$  427.1299; found 427.1296.

The UPCC chromatograms of racemic product **C23**

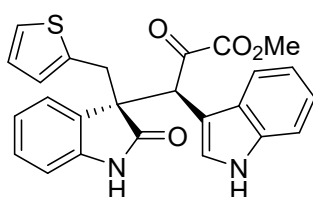


|   | Retention Time | Area     | % Area | Height  |
|---|----------------|----------|--------|---------|
| 1 | 8.157          | 28052570 | 40.60  | 1003896 |
| 2 | 10.068         | 6603605  | 9.56   | 188719  |
| 3 | 15.284         | 27882627 | 40.35  | 557801  |
| 4 | 17.965         | 6560226  | 9.49   | 132875  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 8.179          | 1344302  | 5.04   | 54279  |
| 2 | 15.103         | 25341031 | 94.96  | 484073 |



**C24**

**Methyl (S)-3-(1H-indol-3-yl)-2-oxo-3-((S)-2-oxo-3-(thiophen-2-ylmethyl)indolin-3-yl)propanoate (C24)**

Yellow oil; 35.1 mg, 78% yield, >19/1 dr, 79% ee;  $[\alpha]_D^{20.2} = -149.38$  ( $c = 0.48$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL IG-3,  $\text{CO}_2/\text{MeOH} = 87/13$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 34.76 min,  $t_R$  (minor) = 13.52 min.

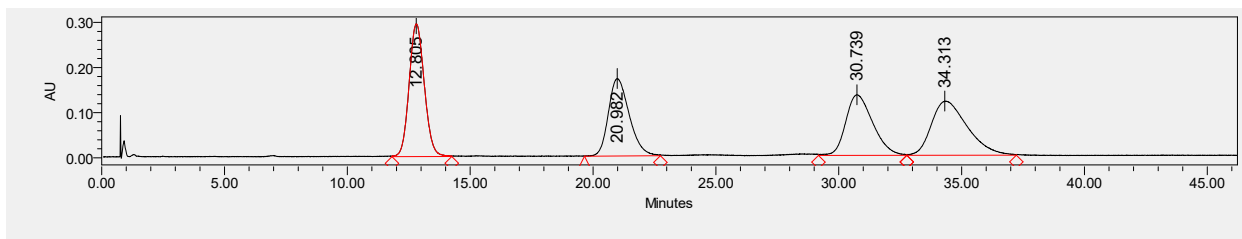
**$^1\text{H}$  NMR** (400 MHz, Chloroform- $d$ )  $\delta$  8.57 (s, 1H), 7.82 (d,  $J = 7.8$  Hz, 1H), 7.66 (s, 1H), 7.46 (d,  $J = 8.0$  Hz, 1H), 7.32 – 7.27 (m, 1H), 7.25 – 7.22 (m, 1H), 7.17 (td,  $J = 7.6, 1.2$  Hz, 1H), 6.92 – 6.87 (m, 2H), 6.85 (dd,  $J = 5.1, 1.2$  Hz, 1H), 6.72 (d,  $J = 8.0$  Hz, 2H), 6.64 (dd,  $J = 5.2, 3.6$  Hz, 1H), 6.49 (d,  $J = 3.2$  Hz, 1H), 5.66 (s, 1H), 3.71 (d,  $J = 14.0$  Hz, 1H), 3.56 (s, 3H), 3.27 (d,  $J = 14.0$  Hz, 1H).

**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CDCl}_3$ )  $\delta$  188.9, 180.0, 161.1, 142.3, 136.7, 136.0, 129.3, 128.5, 127.9, 127.4, 126.5, 126.2, 125.7, 124.7, 123.2, 121.6, 121.0, 119.6, 111.8, 109.8, 105.0, 55.5, 53.0, 51.2, 38.1.

IR: 3370, 2924, 2406, 1727, 1706, 1620, 1470, 1339, 1261, 746  $\text{cm}^{-1}$ .

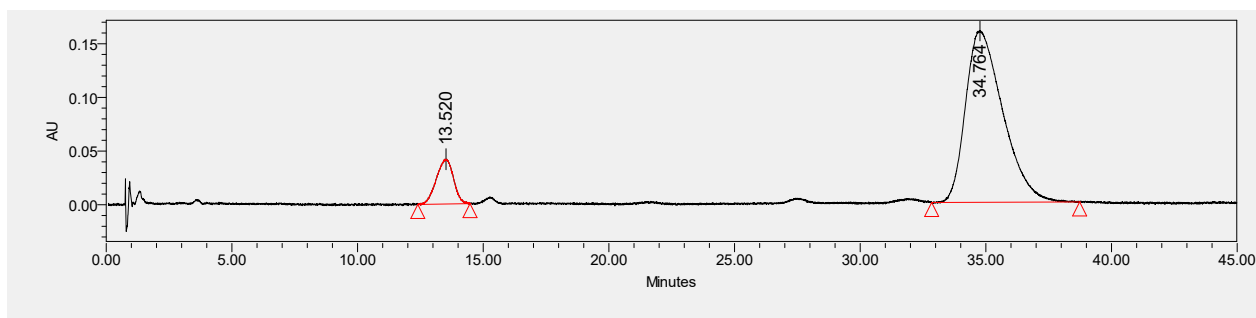
**HRMS** (FTMS+c ESI)  $m/z$ :  $[M - H]^-$  calcd for  $C_{25}H_{20}N_2O_4S$  443.1071; found 443.1070.

The UPCC chromatograms of racemic product **C24**

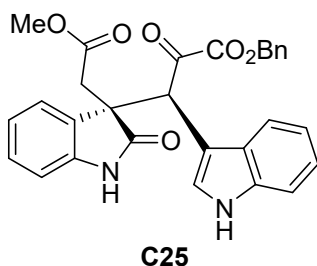


|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 12.805         | 12615645 | 27.92  | 294282 |
| 2 | 20.982         | 10123362 | 22.40  | 171824 |
| 3 | 30.739         | 10240063 | 22.66  | 134301 |
| 4 | 34.313         | 12208997 | 27.02  | 120089 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 13.520         | 1965954  | 10.64  | 41813  |
| 2 | 34.764         | 16516533 | 89.36  | 160175 |



**Benzyl (S)-3-(1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (C25)**

Yellow oil; 35.7 mg, 70% yield, >19:1 dr, 95% ee;  $[\alpha]_D^{19.3} = -224.08$  ( $c = 0.71$  in  $CH_2Cl_2$ ).

**UPCC** DAICEL CHIRALCEL IC-3,  $CO_2/MeOH = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 7.64 min,  $t_R$  (minor) = 6.85 min.

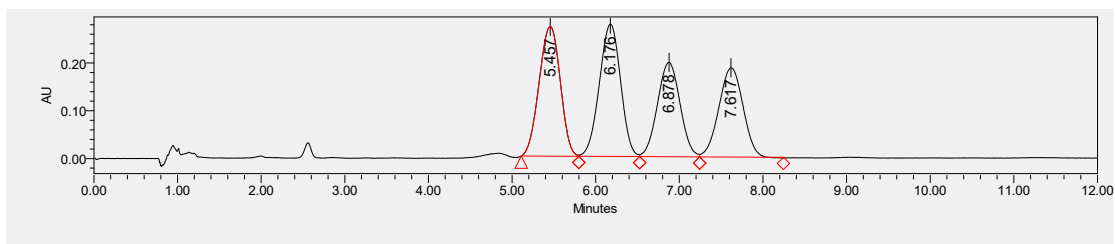
**$^1H$  NMR** (400 MHz,  $CHCl_3-d$ )  $\delta$  8.73 (s, 1H), 8.48 (s, 1H), 7.61 (d,  $J = 8.0$  Hz, 1H), 7.34 (d,  $J = 8.0$  Hz, 1H), 7.25 – 7.14 (m, 4H), 7.11 (t,  $J = 7.6$  Hz, 2H), 6.87 – 6.81 (m, 4H), 6.71 (d,  $J = 8.0$  Hz, 2H), 5.45 (s, 1H), 4.92 (q,  $J = 12.4$  Hz, 2H), 3.32 (s, 3H), 3.13 (dd,  $J = 56.0, 16.0$  Hz, 2H).

**$^{13}C$  NMR** (101 MHz,  $CDCl_3$ )  $\delta$  187.7, 180.1, 169.9, 160.5, 142.4, 136.6, 134.2, 129.2, 128.7, 128.6, 128.5, 128.5, 128.2, 127.8, 126.9, 125.0, 123.0, 121.7, 120.9, 119.1, 111.9, 110.1, 103.8, 68.0, 51.7, 51.3, 50.5, 40.4.

**IR**: 3367, 2923, 2351, 1728, 1620, 1470, 1340, 1261, 1210, 746  $cm^{-1}$ .

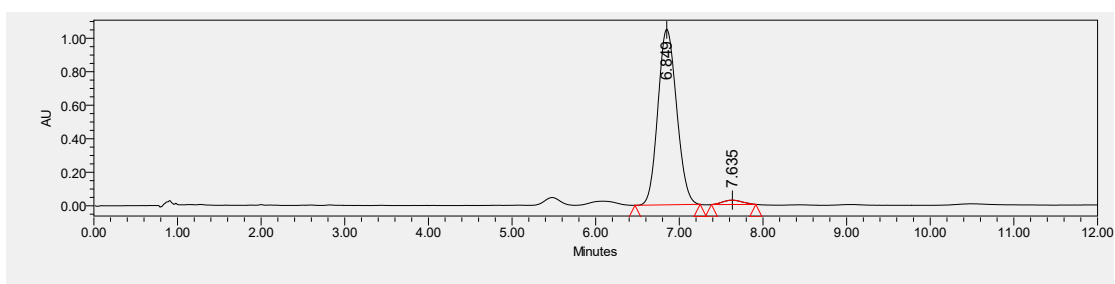
**HRMS** (FTMS+c ESI)  $m/z$ :  $[M - H]^-$  calcd for  $C_{30}H_{26}N_2O_6$  509.1718.

The UPCC chromatograms of racemic product **C25**

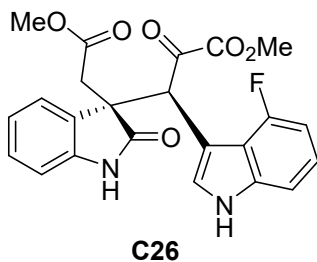


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 5.457          | 4761008 | 27.88  | 271524 |
| 2 | 6.176          | 4873949 | 28.54  | 276895 |
| 3 | 6.878          | 3691720 | 21.62  | 197259 |
| 4 | 7.617          | 3750094 | 21.96  | 186672 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height  |
|---|----------------|----------|--------|---------|
| 1 | 6.849          | 16278429 | 97.47  | 1048542 |
| 2 | 7.635          | 422978   | 2.53   | 26018   |



**Methyl (S)-3-(4-fluoro-1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (C26)**

Yellow oil; 39.0 mg, 89% yield, >19:1 dr, 93% ee;  $[\alpha]_D^{19.5} = -245.55$  ( $c = 0.29$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL OD-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 2.49 min,  $t_R$  (minor) = 3.10 min.

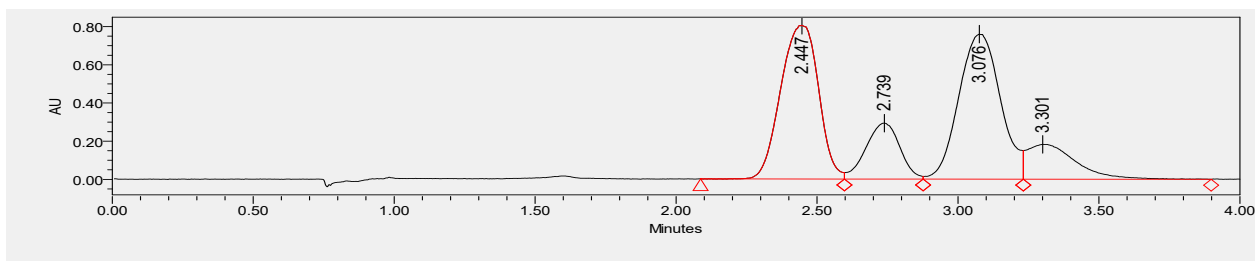
**$^1\text{H}$  NMR** (400 MHz, Chloroform- $d$ )  $\delta$  8.99 (s, 1H), 8.31 (s, 1H), 7.20 (td,  $J = 7.6, 1.2$  Hz, 1H), 7.15 – 7.06 (m, 2H), 6.96 – 6.78 (m, 3H), 6.78 – 6.69 (m, 1H), 6.66 (d,  $J = 2.8$  Hz, 1H), 5.69 (d,  $J = 1.6$  Hz, 1H), 3.59 (s, 3H), 3.34 – 3.30 (m, 4H), 3.14 (d,  $J = 15.9$  Hz, 1H).

**$^{13}\text{C}$  NMR** (101 MHz, Chloroform- $d$ )  $\delta$  187.76, 179.98, 170.12, 160.87, 156.97 (d,  $J = 247.1$  Hz), 142.34, 139.07 (d,  $J = 10.2$  Hz), 129.24, 128.72, 126.92, 125.04, 123.63 (d,  $J = 8.1$  Hz), 121.71, 116.70 (d,  $J = 18.1$  Hz), 110.01, 108.19 (d,  $J = 3.5$  Hz), 106.17 (d,  $J = 19.1$  Hz), 102.28, 53.23, 51.70, 51.40, 50.44, 39.53.

**IR**: 3352, 2954, 2350, 1731, 1621, 1470, 1346, 1230, 1034, 742  $\text{cm}^{-1}$ .

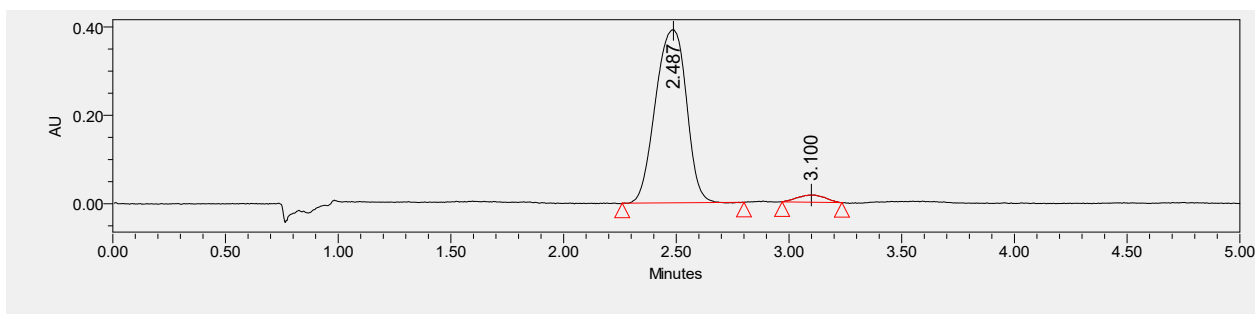
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{23}\text{H}_{19}\text{FN}_2\text{O}_6$  437.1154; found 437.1152.

The UPCC chromatograms of racemic product **C26**

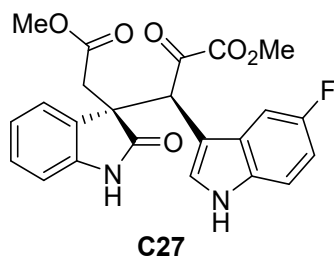


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 2.447          | 7508697 | 37.60  | 804623 |
| 2 | 2.739          | 2451071 | 12.27  | 292639 |
| 3 | 3.076          | 7805039 | 39.08  | 759438 |
| 4 | 3.301          | 2205000 | 11.04  | 182583 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 2.487          | 3742058 | 96.44  | 392431 |
| 2 | 3.100          | 138226  | 3.56   | 16253  |



**Methyl (S)-3-(5-fluoro-1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (C27)**

Yellow oil; 38.1 mg, 87% yield, >19:1 dr, 96% ee;  $[\alpha]_D^{19.7} = -264.28$  ( $c = 0.45$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL OZ-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 5.08 min,  $t_R$  (minor) = 6.81 min.

**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.92 (d,  $J = 2.8$  Hz, 1H), 8.53 (s, 1H), 7.30 – 7.16 (m, 3H), 6.93 (td,  $J = 9.2, 2.4$  Hz, 1H), 6.90 – 6.83 (m, 3H), 6.77 (d,  $J = 7.6$  Hz, 1H), 5.39 (s, 1H), 3.57 (s, 3H), 3.36 (s, 3H), 3.15 ( $J = 28.0, 16.0$  Hz, 2H).

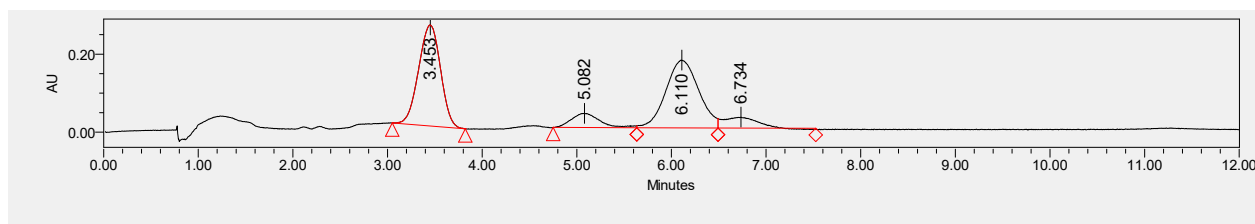
**$^{13}\text{C}$  NMR** (101 MHz,  $\text{Chloroform-}d$ )  $\delta$  187.72, 179.97, 170.05, 160.88, 158.51 (d,  $J = 215.2$  Hz), 142.15, 132.98, 129.38, 128.86, 128.52, 128.27 (d,  $J = 9.6$  Hz), 124.85, 121.89, 112.72 (d,  $J = 9.7$  Hz), 111.61 (d,  $J = 26.6$  Hz), 110.16, 104.04 (d,  $J = 1.8$  Hz), 103.90 (d,  $J = 17.2$  Hz), 53.24, 51.83, 51.55, 49.86, 39.83.

**$^{19}\text{F}$  NMR** (377 MHz,  $\text{CDCl}_3$ )  $\delta$  -122.6.

**IR**: 3353, 2954, 2350, 1728, 1621, 1486, 1344, 1259, 1167, 753  $\text{cm}^{-1}$ .

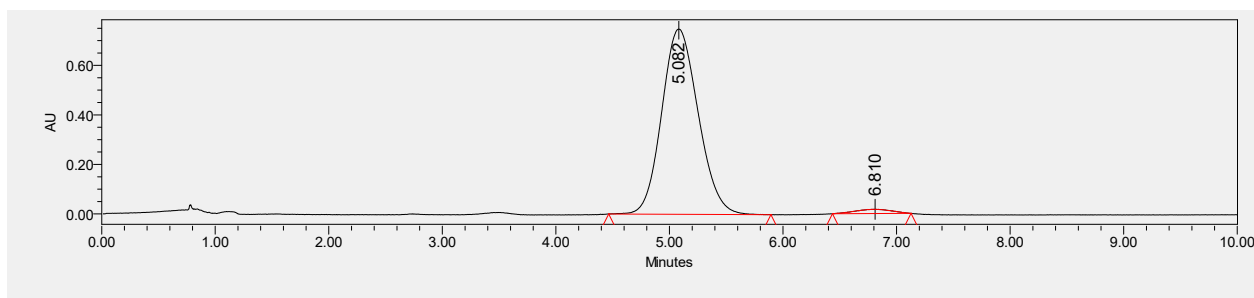
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{23}\text{H}_{19}\text{FN}_2\text{O}_6$  437.1154; found 437.1153.

The UPCC chromatograms of racemic product **C27**

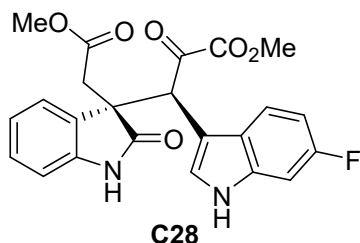


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 3.453          | 4151127 | 42.22  | 259100 |
| 2 | 5.082          | 760341  | 7.73   | 36273  |
| 3 | 6.110          | 4182120 | 42.54  | 173986 |
| 4 | 6.734          | 738110  | 7.51   | 27966  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 5.082          | 16709597 | 97.79  | 748047 |
| 2 | 6.810          | 377912   | 2.21   | 17143  |



**Methyl (S)-3-(6-fluoro-1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (C28)**

Yellow solid; 41.6 mg, 95% yield, >19:1 dr, 95% ee; melting point: 108 – 110 °C;  $[\alpha]_D^{19.0} = -219.23$  ( $c = 0.73$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL OX-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 5.03 min,  $t_R$  (minor) = 8.08 min.

**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.97 (s, 1H), 8.68 (s, 1H), 7.51 (dd,  $J = 8.8, 5.2$  Hz, 1H), 7.19 (t,  $J = 7.6$  Hz, 1H), 7.00 (dd,  $J = 9.2, 2.4$  Hz, 1H), 6.95 – 6.73 (m, 5H), 5.44 (s, 1H), 3.56 (s, 3H), 3.35 (s, 3H), 3.15 (dd,  $J = 24.0, 16.0$  Hz, 2H).

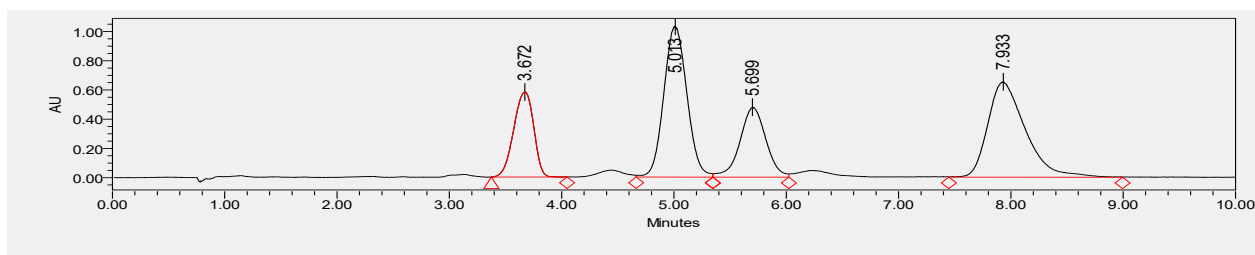
**$^{13}\text{C}$  NMR** (101 MHz,  $\text{Chloroform-}d$ )  $\delta$  187.72, 180.16, 170.09, 160.93, 160.26 (d,  $J = 12.4$  Hz), 142.16, 136.49 (d,  $J = 12.4$  Hz), 129.39, 128.86, 127.23, 124.86, 124.25, 121.88, 119.82 (d,  $J = 10.1$  Hz), 110.20, 109.73 (d,  $J = 24.8$  Hz), 104.00, 98.21 (d,  $J = 26.2$  Hz), 53.24, 51.83, 51.50, 49.78, 39.83.

**$^{19}\text{F}$  NMR** (377 MHz,  $\text{CDCl}_3$ )  $\delta$  -119.8.

**IR**: 3355, 2954, 2350, 1730, 1623, 1471, 1343, 1261, 1143, 753  $\text{cm}^{-1}$ .

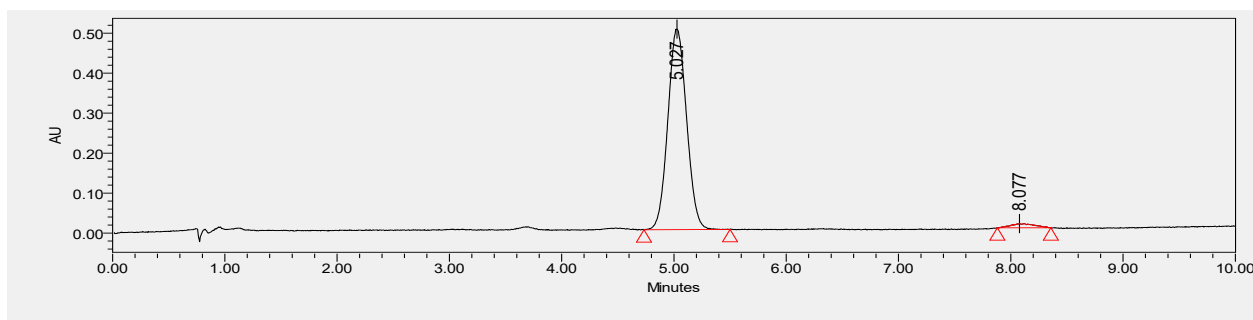
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{23}\text{H}_{19}\text{FN}_2\text{O}_6$  437.1154; found 437.1154.

The UPCC chromatograms of racemic product **C28**

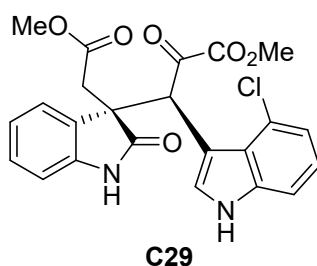


|   | Retention Time | Area     | % Area | Height  |
|---|----------------|----------|--------|---------|
| 1 | 3.672          | 7197044  | 16.09  | 582300  |
| 2 | 5.013          | 14769920 | 33.03  | 1032307 |
| 3 | 5.699          | 7631124  | 17.06  | 478656  |
| 4 | 7.933          | 15124150 | 33.82  | 652783  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 5.027          | 5825617 | 97.43  | 501866 |
| 2 | 8.077          | 153513  | 2.57   | 10571  |



**Methyl (S)-3-(4-chloro-1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (29)**

Yellow oil; 37.7 mg, 83% yield, >19:1 dr, 90% ee;  $[\alpha]_D^{19.3} = -338.38$  ( $c = 0.28$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL IC-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 4.43 min,  $t_R$  (minor) = 6.34 min.

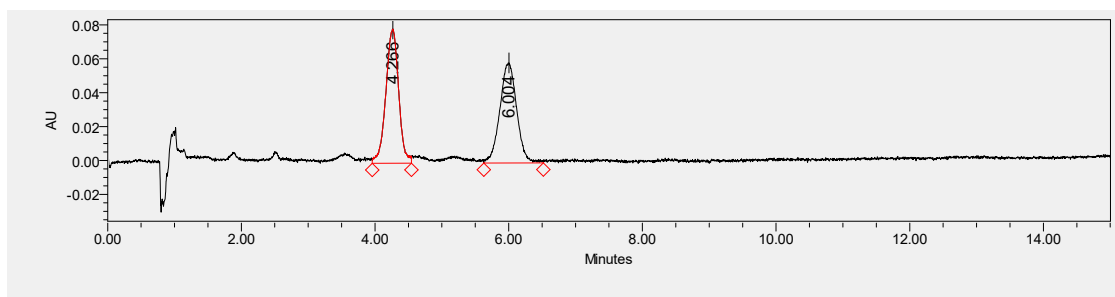
**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  9.30 (d,  $J = 2.8$  Hz, 1H), 8.44 (s, 1H), 7.23-7.13 (m, 3H), 7.06 (t,  $J = 7.6$  Hz, 1H), 6.92 – 6.84 (m, 2H), 6.80 – 6.74 (m, 1H), 6.67 (d,  $J = 2.8$  Hz, 1H), 6.25 (s, 1H), 3.57 (s, 3H), 3.29 (dd,  $J = 96.0, 16.0$  Hz) 3.27 (s, 3H).

**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CDCl}_3$ )  $\delta$  187.6, 180.2, 170.3, 161.0, 142.4, 138.1, 129.5, 128.7, 128.6, 126.0, 125.3, 123.8, 123.5, 122.3, 121.8, 111.2, 110.1, 103.4, 53.2, 51.7, 51.3, 49.3, 39.4.

**IR**: 3349, 2921, 2350, 1730, 1620, 1470, 1340, 1261, 1117, 747  $\text{cm}^{-1}$ .

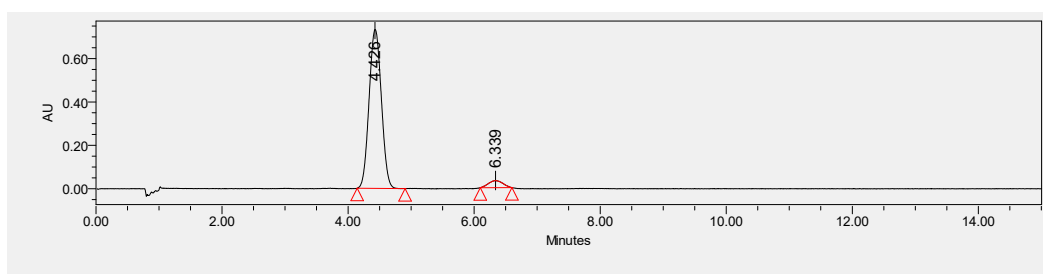
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{23}\text{H}_{19}\text{ClN}_2\text{O}_6$  453.0859, 455.0829; found 453.0858, 455.0828.

The UPCC chromatograms of racemic product **C29**

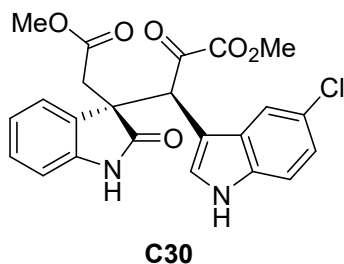


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 4.266          | 1049949 | 50.38  | 79208  |
| 2 | 6.004          | 1034051 | 49.62  | 59073  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 4.426          | 9889274 | 94.95  | 732609 |
| 2 | 6.339          | 525524  | 5.05   | 33125  |



**Methyl (S)-3-(5-chloro-1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (C30)**

Yellow oil; 39.1 mg, 86% yield, >19:1 dr, 91% ee;  $[\alpha]_D^{19.3} = -197.86$  ( $c = 0.28$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL OZ-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 6.22 min,  $t_R$  (minor) = 8.27 min.

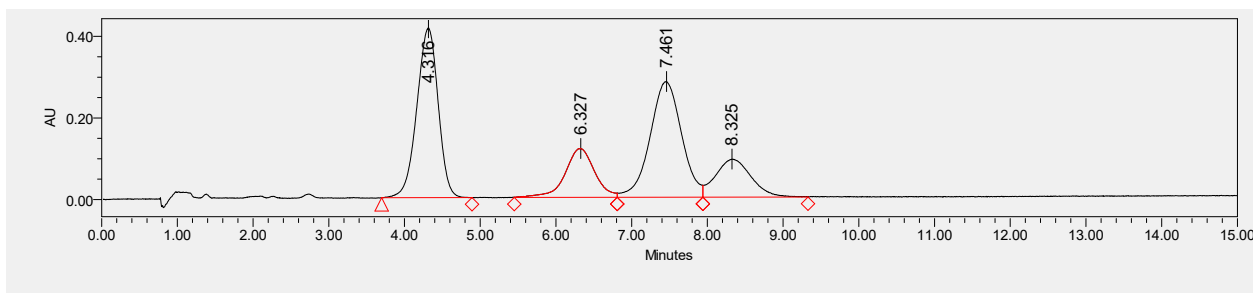
**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.95 (s, 1H), 8.55 (s, 1H), 7.59 (d,  $J = 1.9$  Hz, 1H), 7.25 – 7.11 (m, 3H), 6.92 – 6.77 (m, 4H), 5.41 (s, 1H), 3.58 (s, 3H), 3.36 (s, 3H), 3.16 (dd,  $J = 24.0, 16.0$  Hz, 2H).

**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CDCl}_3$ )  $\delta$  187.7, 180.0, 170.1, 160.9, 142.1, 134.8, 129.4, 128.9, 128.9, 128.1, 126.6, 124.8, 123.5, 121.9, 118.4, 113.0, 110.2, 103.8, 53.3, 51.9, 51.5, 49.6, 39.7.

**IR**: 3352, 2925, 2350, 1730, 1620, 1469, 1342, 1260, 1082, 747  $\text{cm}^{-1}$ .

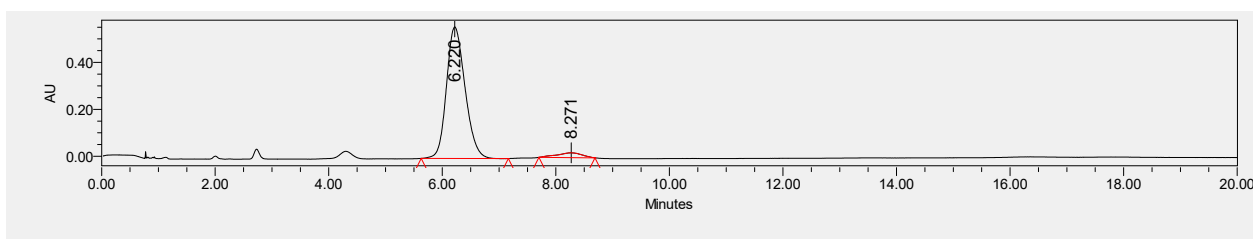
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{23}\text{H}_{19}\text{ClN}_2\text{O}_6$  453.0859, 455.0829; found 453.0858, 455.0830.

The UPCC chromatograms of racemic product **C30**

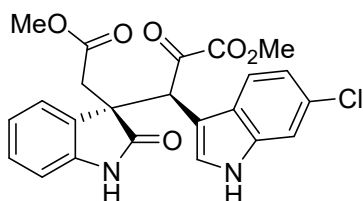


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 4.316          | 8111396 | 35.94  | 416322 |
| 2 | 6.327          | 3232475 | 14.32  | 119650 |
| 3 | 7.461          | 8198600 | 36.33  | 283675 |
| 4 | 8.325          | 3026580 | 13.41  | 93076  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 6.220          | 12741402 | 95.31  | 560657 |
| 2 | 8.271          | 626557   | 4.69   | 20544  |



**C31**

**Methyl (S)-3-(6-chloro-1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (C31)**

Yellow oil; 35.0 mg, 77% yield, >19:1 dr, 94% ee;  $[\alpha]_D^{19.3} = -197.86$  ( $c = 0.17$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL OX-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 6.91 min,  $t_R$  (minor) = 12.40 min.

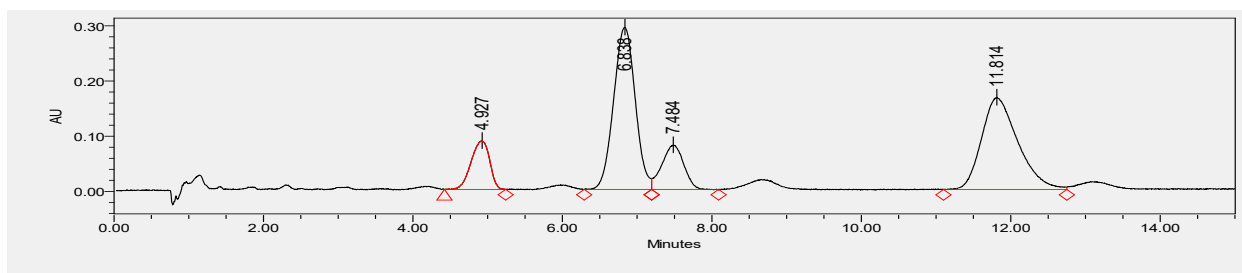
**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.85 (s, 1H), 8.49 (s, 1H), 7.50 (d,  $J = 8.6$  Hz, 1H), 7.33 (d,  $J = 1.6$  Hz, 1H), 7.19 (td,  $J = 7.6, 1.2$  Hz, 1H), 7.11 (dd,  $J = 8.8, 1.6$  Hz, 1H), 6.91 – 6.82 (m, 3H), 6.76 (d,  $J = 7.6$  Hz, 1H), 5.44 (s, 1H), 3.57 (s, 3H), 3.37 (s, 3H), 3.14 (dd,  $J = 24.0, 12.0$  Hz, 2H).

**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CDCl}_3$ )  $\delta$  187.8, 180.0, 170.0, 160.9, 142.1, 136.8, 129.4, 128.9, 127.4, 126.3, 124.8, 121.9, 121.7, 119.9, 111.8, 110.2, 104.3, 53.3, 51.9, 51.5, 49.7, 39.8.

**IR**: 3365, 2955, 2350, 1731, 1620, 1471, 1338, 1260, 1116, 753  $\text{cm}^{-1}$ .

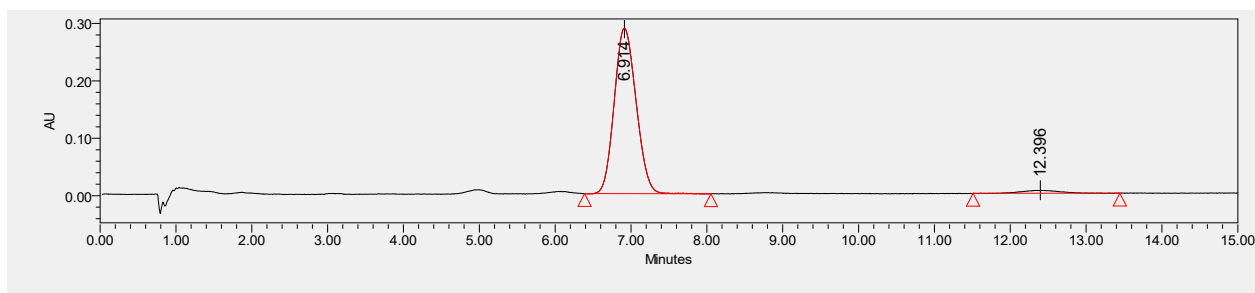
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{23}\text{H}_{19}\text{ClN}_2\text{O}_6$  453.0859, 455.0829; found 453.0856, 455.0829.

The UPCC chromatograms of racemic product **C31**

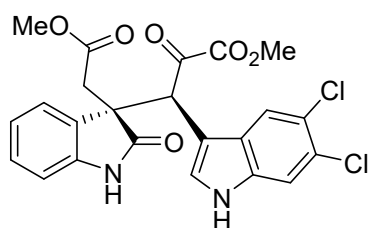


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 4.927          | 1518670 | 10.34  | 88574  |
| 2 | 6.838          | 5901286 | 40.20  | 294153 |
| 3 | 7.484          | 1638382 | 11.16  | 81003  |
| 4 | 11.814         | 5622979 | 38.30  | 167003 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 6.914          | 5764276 | 97.04  | 288026 |
| 2 | 12.396         | 175775  | 2.96   | 5231   |



**C32**

**Methyl (S)-3-(5,6-dichloro-1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (C32)**

Yellow solid; 39.9 mg, 80% yield, >19:1 dr, 97% ee; melting point: 117 – 123 °C;  $[\alpha]_D^{20.1} = -204.17$  ( $c = 0.24$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL OD-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 4.37 min,  $t_R$  (minor) = 3.96 min.

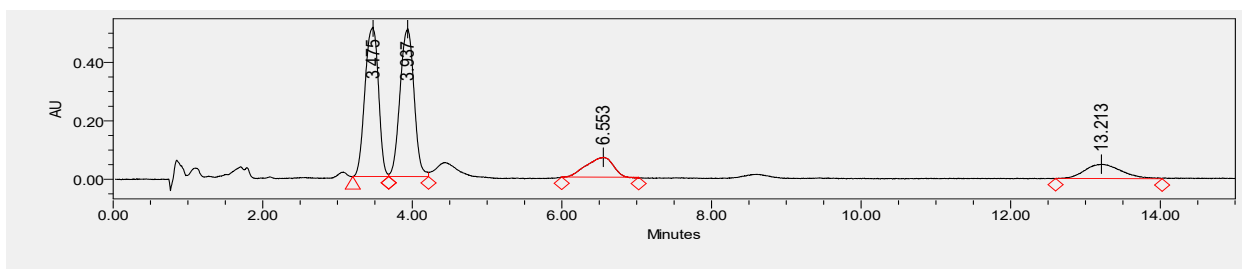
**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-d}$ )  $\delta$  9.10 (s, 1H), 8.66 (s, 1H), 7.64 (s, 1H), 7.42 (s, 1H), 7.19 (td,  $J = 7.6, 1.4$  Hz, 1H), 6.96 – 6.82 (m, 3H), 6.77 (d,  $J = 7.6$  Hz, 1H), 5.38 (s, 1H), 3.60 (s, 3H), 3.40 (s, 3H), 3.20 (dd,  $J = 32.0, 12.0$  Hz, 2H).

**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CDCl}_3$ )  $\delta$  187.7, 179.9, 170.2, 160.8, 141.9, 135.2, 129.5, 129.1, 128.5, 127.5, 126.9, 125.0, 124.6, 122.1, 119.9, 113.5, 110.3, 103.9, 53.4, 52.0, 51.6, 49.4, 39.0.

**IR**: 3341, 2917, 2350, 1731, 1621, 1471, 1451, 1229, 1082, 753  $\text{cm}^{-1}$ .

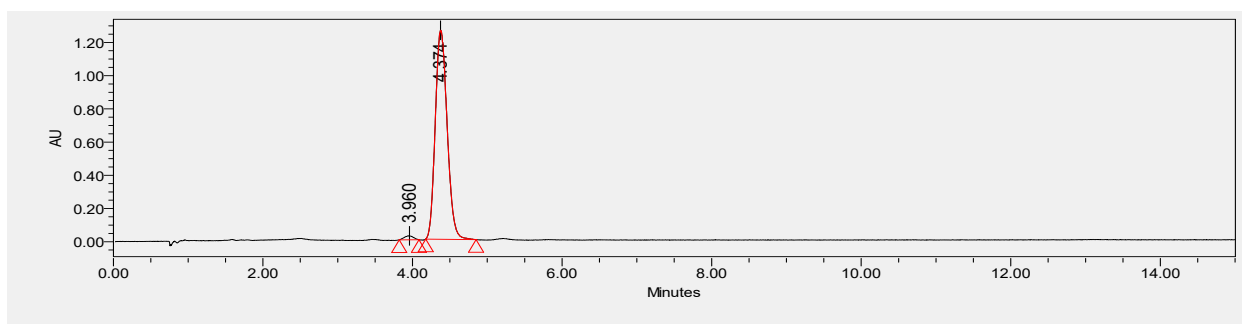
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{23}\text{H}_{18}\text{Cl}_2\text{N}_2\text{O}_6$  487.0469, 489.0440; found 487.0468, 489.0440.

The UPCC chromatograms of racemic product **C32**

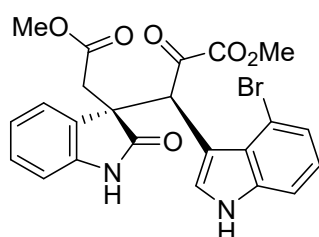


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 3.475          | 6397951 | 39.19  | 511796 |
| 2 | 3.937          | 6581968 | 40.31  | 507075 |
| 3 | 6.553          | 1693434 | 10.37  | 67883  |
| 4 | 13.213         | 1653446 | 10.13  | 49396  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height  |
|---|----------------|----------|--------|---------|
| 1 | 3.960          | 204646   | 1.45   | 24006   |
| 2 | 4.374          | 13862607 | 98.55  | 1258819 |



**C33**

**Methyl (S)-3-(4-bromo-1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (C33)**

Yellow oil; 47.3 mg, 95% yield, >19:1 dr, 84% ee;  $[\alpha]_D^{19.1} = -259.91$  ( $c = 0.24$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL AD-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 6.61 min,  $t_R$  (minor) = 5.47 min.

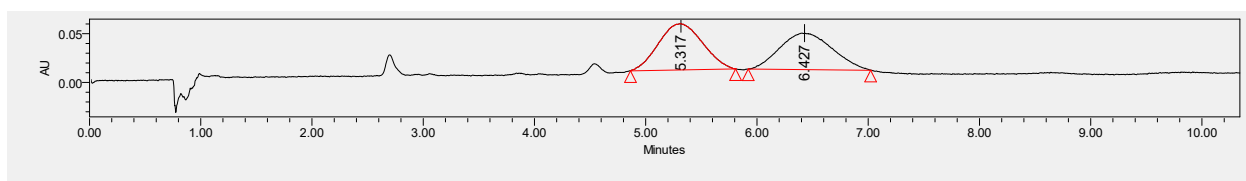
**$^1\text{H}$  NMR** (400 MHz, Acetonitrile- $d_3$ )  $\delta$  9.90 (s, 1H), 8.66 (s, 1H), 7.52 (dd,  $J = 8.4, 0.8$  Hz, 1H), 7.38 (dd,  $J = 7.6, 0.8$  Hz, 1H), 7.24 (td,  $J = 7.6, 1.2$  Hz, 1H), 7.11 (t,  $J = 7.8$  Hz, 1H), 6.97 – 6.87 (m, 3H), 6.81 (d,  $J = 7.6$  Hz, 1H), 6.26 (s, 1H), 3.52 (s, 3H), 3.41 (d,  $J = 15.6$  Hz, 1H), 3.24 (s, 3H), 2.92 (d,  $J = 15.6$  Hz, 1H).

**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CD}_3\text{CN}$ )  $\delta$  188.9, 180.1, 170.6, 162.2, 144.3, 139.2, 130.5, 130.4, 129.3, 126.3, 126.2, 125.6, 124.4, 122.0, 114.1, 113.0, 110.3, 104.5, 53.5, 51.9, 51.7, 49.6, 40.5.

**IR**: 3348, 2921, 2350, 1729, 1620, 1437, 1337, 1176, 1080, 745  $\text{cm}^{-1}$ .

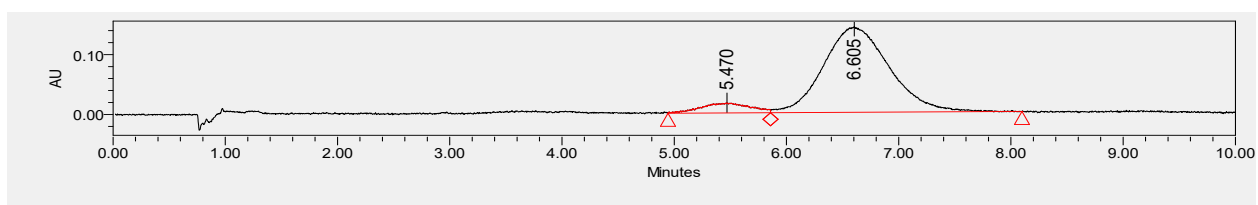
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{23}\text{H}_{19}\text{BrN}_2\text{O}_6$  497.0354, 499.0334; found 497.0352, 499.0333.

The UPCC chromatograms of racemic product **C33**

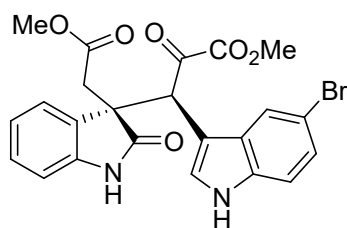


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 5.317          | 1298845 | 51.42  | 47429  |
| 2 | 6.427          | 1226904 | 48.58  | 37440  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 5.470          | 502802  | 7.99   | 16907  |
| 2 | 6.605          | 5790698 | 92.01  | 143386 |



**C34**

**Methyl (S)-3-(5-bromo-1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (C34)**

Yellow solid; 41.0 mg, 84% yield, >19:1 dr, 97% ee; melting point: 127 – 133 °C;  $[\alpha]_D^{20.5} = -252.35$  ( $c = 0.43$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL OZ-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 7.36 min,  $t_R$  (minor) = 5.08 min.

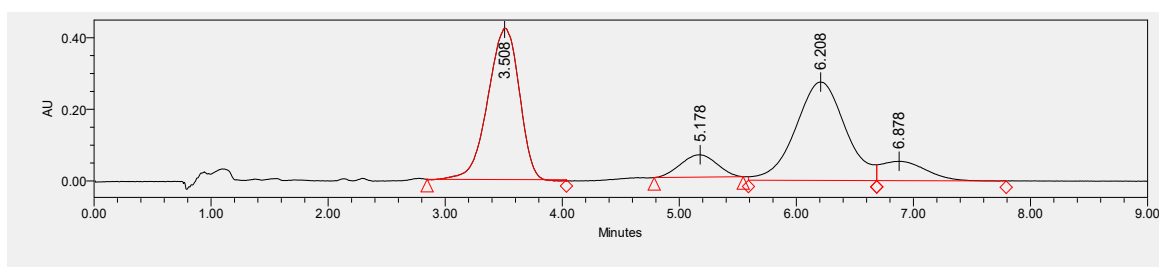
**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.96 (s, 1H), 8.54 (s, 1H), 7.74 (s, 1H), 7.28 – 7.24 (m, 1H), 7.23 – 7.16 (m, 2H), 6.89 (t,  $J = 7.3$  Hz, 1H), 6.85 – 6.79 (m, 3H), 5.41 (s, 1H), 3.58 (s, 3H), 3.36 (s, 3H), 3.16 (dd,  $J = 20.0, 16.0$  Hz, 2H).

**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CDCl}_3$ )  $\delta$  187.7, 179.9, 170.1, 160.9, 142.1, 135.1, 129.5, 129.4, 128.9, 127.9, 126.0, 124.8, 122.0, 121.5, 114.2, 113.4, 110.2, 103.7, 53.3, 51.9, 51.5, 49.6, 39.6.

**IR**: 3363, 2920, 2350, 1730, 1620, 1469, 1259, 1115, 1080, 753  $\text{cm}^{-1}$ .

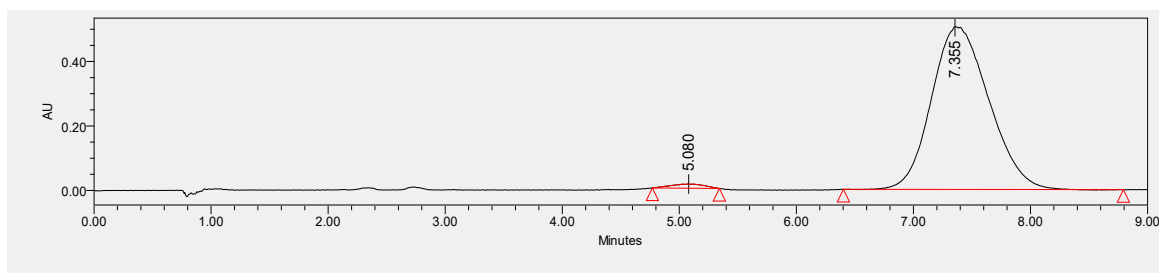
**HRMS** (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{23}\text{H}_{19}\text{BrN}_2\text{O}_6$  497.0354, 499.0334; found 497.0344, 499.0332.

The UPCC chromatograms of racemic product **C34**

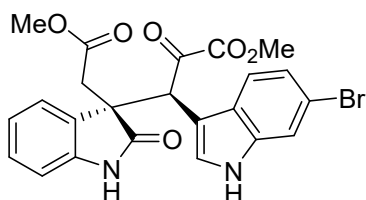


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 3.508          | 7890874 | 42.00  | 422518 |
| 2 | 5.178          | 1347142 | 7.17   | 62373  |
| 3 | 6.208          | 8116195 | 43.20  | 274524 |
| 4 | 6.878          | 1431702 | 7.62   | 53885  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 5.080          | 258606   | 1.48   | 13228  |
| 2 | 7.355          | 17221877 | 98.52  | 504941 |



**C35**

**Methyl (S)-3-(6-bromo-1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (C35)**

Yellow solid; 41.0 mg, 92% yield, 92:8 dr, 95/81% ee; melting point: 135 – 136 °C;  $[\alpha]_D^{20.6} = -$

166.97 (*c* = 0.25 in CH<sub>2</sub>Cl<sub>2</sub>).

**UPCC** DAICEL CHIRALCEL OX-3, CO<sub>2</sub>/MeOH = 80/20, flow rate = 1.5 mL/min,  $\lambda$  = 211 nm, major isomer: *t<sub>R</sub>* (major) = 8.26 min, *t<sub>R</sub>* (minor) = 14.70 min; minor isomer: *t<sub>R</sub>* (major) = 5.82 min, *t<sub>R</sub>* (minor) = 9.12 min.

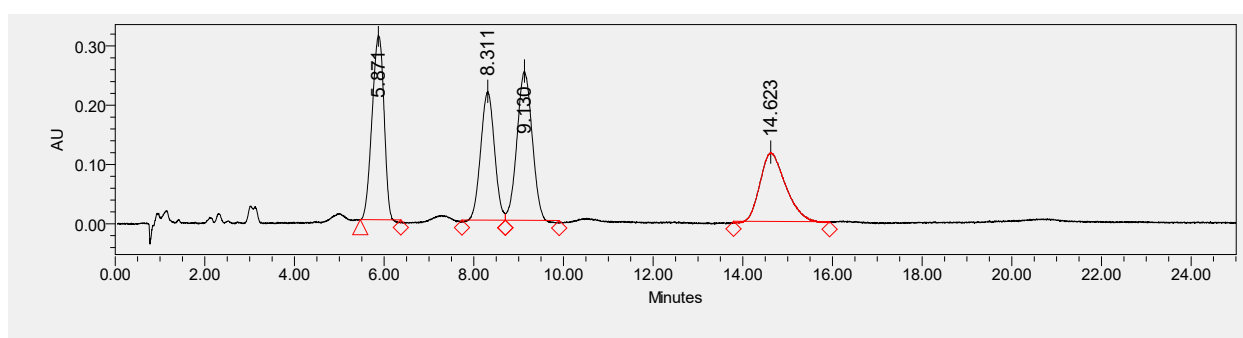
**<sup>1</sup>H NMR** (400 MHz, Chloroform-*d*)  $\delta$  9.04 (s, 1H), 8.77 (s, 1H), 7.48 (s, 1H), 7.43 (d, *J* = 8.5 Hz, 1H), 7.19 (dd, *J* = 16.4, 8.4 Hz, 2H), 6.89 – 6.77 (m, 4H), 5.43 (s, 1H), 3.55 (s, 3H), 3.36 (s, 3H), 3.16 (t, *J* = 16.0, 2H).

**<sup>13</sup>C NMR** (101 MHz, CDCl<sub>3</sub>)  $\delta$  187.8, 180.1, 170.1, 160.9, 142.1, 137.2, 129.4, 128.9, 127.5, 126.6, 124.8, 124.1, 122.0, 120.2, 116.5, 114.9, 110.3, 104.2, 53.3, 51.9, 51.5, 49.7, 39.6.

**IR**: 3353, 2922, 2350, 1730, 1618, 1470, 1259, 1115, 1052, 754 cm<sup>-1</sup>.

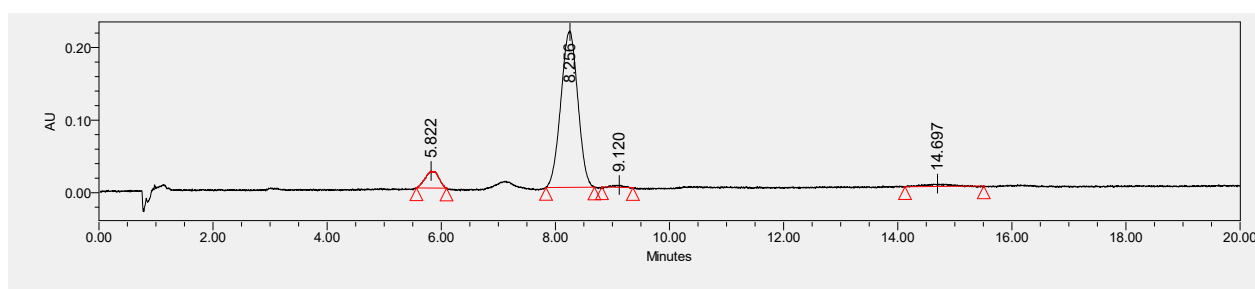
**HRMS** (FTMS+c ESI) *m/z*: [M - H]<sup>-</sup> calcd for C<sub>23</sub>H<sub>19</sub>BrN<sub>2</sub>O<sub>6</sub> 497.0354, 499.0334; found 497.0353, 499.0334.

The UPCC chromatograms of racemic product **C35**

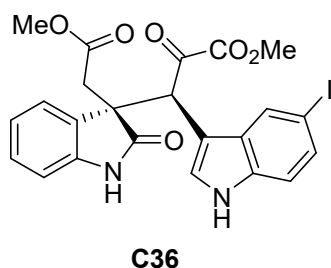


| f | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 5.871          | 5762791 | 27.52  | 311375 |
| 2 | 8.311          | 4639510 | 22.16  | 217601 |
| 3 | 9.130          | 5959399 | 28.46  | 251828 |
| 4 | 14.623         | 4576815 | 21.86  | 116971 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 5.822          | 383217  | 7.72   | 23479  |
| 2 | 8.256          | 4425303 | 89.14  | 215310 |
| 3 | 9.120          | 40909   | 0.82   | 3207   |
| 4 | 14.697         | 114770  | 2.31   | 4025   |



**Methyl (S)-3-(5-iodo-1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (C36)**

Yellow solid; 49.7 mg, 91% yield, 88:12 dr, 94/80% ee; melting point: 153-156 °C; [ $\alpha$ ]<sub>D</sub><sup>19.8</sup> = -165.01

(*c* = 0.3 in CH<sub>2</sub>Cl<sub>2</sub>).

**UPCC** DAICEL CHIRALCEL OX-3, CO<sub>2</sub>/MeOH = 80/20, flow rate = 1.5 mL/min,  $\lambda$  = 211 nm, major isomer: *t<sub>R</sub>* (major) = 9.03 min, *t<sub>R</sub>* (minor) = 12.09 min; minor isomer: *t<sub>R</sub>* (major) = 6.29 min, *t<sub>R</sub>* (minor) = 10.11 min.

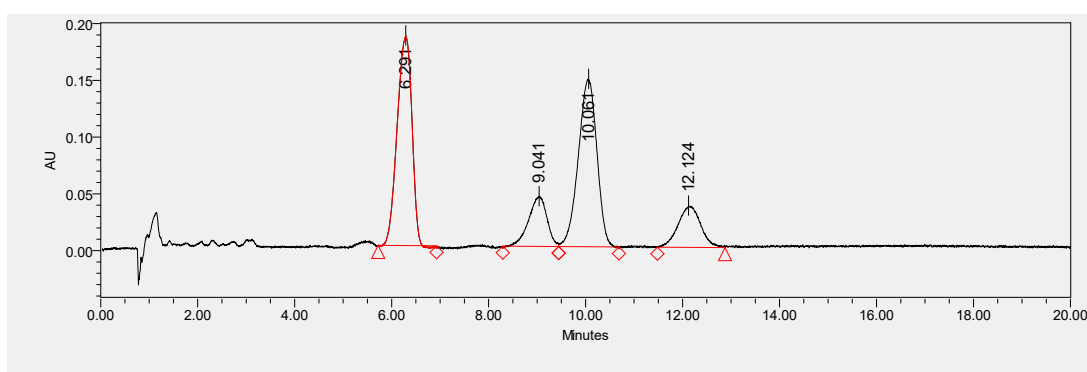
**<sup>1</sup>H NMR** (400 MHz, Chloroform-*d*)  $\delta$  8.94 (s, 1H), 8.54 (s, 1H), 7.93 (d, *J* = 1.6 Hz, 1H), 7.43 (d, *J* = 8.4 Hz, 1H), 7.21 (t, *J* = 7.6 Hz, 1H), 7.10 (d, *J* = 8.4 Hz, 1H), 6.91 (t, *J* = 7.6 Hz, 1H), 6.85 – 6.78 (m, 3H), 5.41 (s, 1H), 3.58 (s, 3H), 3.37 (s, 3H), 3.14 (dd, *J* = 20.0, 16.0 Hz, 2H).

**<sup>13</sup>C NMR** (101 MHz, CDCl<sub>3</sub>)  $\delta$  187.7, 179.9, 170.1, 160.9, 142.1, 135.6, 131.4, 130.2, 129.4, 129.0, 127.7, 127.4, 124.8, 122.0, 113.8, 110.2, 103.4, 84.4, 53.3, 51.9, 51.6, 49.6, 39.6.

**IR**: 3364, 2921, 2350, 1730, 1620, 1470, 1260, 1115, 1081, 755 cm<sup>-1</sup>.

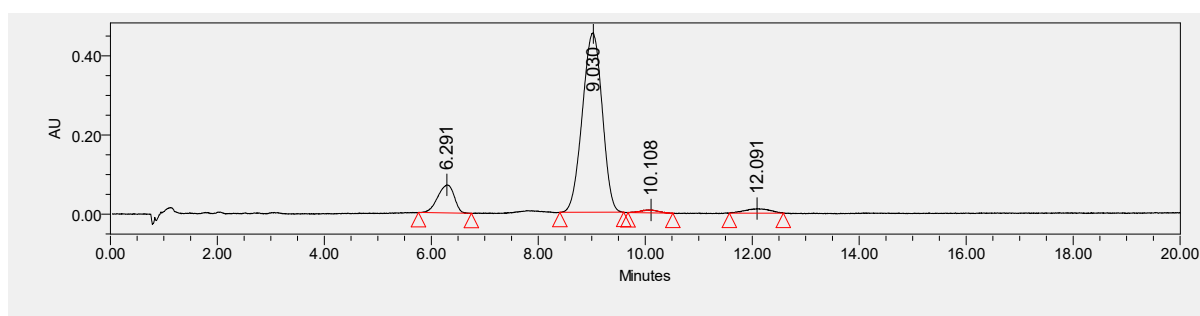
**HRMS** (FTMS+c ESI) *m/z*: [M - H]<sup>-</sup> calcd for C<sub>23</sub>H<sub>19</sub>IN<sub>2</sub>O<sub>6</sub> 545.0215; found 545.0214.

The UPCC chromatograms of racemic product **C36**

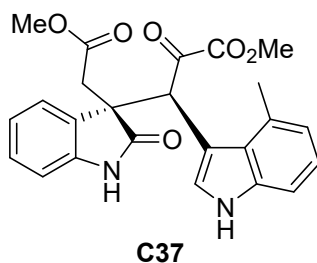


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 6.291          | 3921015 | 38.46  | 185241 |
| 2 | 9.041          | 1160012 | 11.38  | 44147  |
| 3 | 10.061         | 3931082 | 38.56  | 147977 |
| 4 | 12.124         | 1182727 | 11.60  | 36986  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 6.291          | 1503579  | 10.98  | 70799  |
| 2 | 9.030          | 11642446 | 85.06  | 453667 |
| 3 | 10.108         | 169525   | 1.24   | 7582   |
| 4 | 12.091         | 372549   | 2.72   | 11596  |



**Methyl (S)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-3-(4-methyl-1H-indol-3-yl)-2-oxopropanoate (C37)**

Yellow solid; 42.1 mg, 97% yield, >19:1 dr, 91% ee; melting point: 116-121 °C; [ $\alpha$ ]<sub>D</sub><sup>19.5</sup> = -295.95 (*c*

= 0.64 in CH<sub>2</sub>Cl<sub>2</sub>).

**UPCC** DAICEL CHIRALCEL IE-3, CO<sub>2</sub>/MeOH = 80/20, flow rate = 1.5 mL/min,  $\lambda$  = 211 nm,  $t_R$  (major) = 8.05 min,  $t_R$  (minor) = 7.15 min.

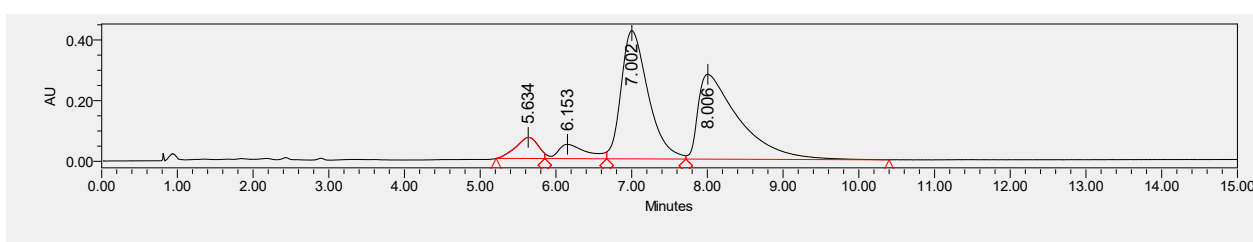
**<sup>1</sup>H NMR** (400 MHz, Chloroform-*d*)  $\delta$  8.86 (d,  $J$  = 2.9 Hz, 1H), 8.49 (s, 1H), 7.25 – 7.20 (m, 1H), 7.13 (d,  $J$  = 8.1 Hz, 1H), 7.07 (dd,  $J$  = 8.4, 7.2 Hz, 1H), 6.93 (d,  $J$  = 7.2 Hz, 1H), 6.87 (d,  $J$  = 7.6 Hz, 2H), 6.85 – 6.80 (m, 1H), 6.69 (d,  $J$  = 2.8 Hz, 1H), 5.74 (s, 1H), 3.57 (s, 3H), 3.34 – 3.14 (m, 5H), 2.87 (s, 3H).

**<sup>13</sup>C NMR** (101 MHz, CDCl<sub>3</sub>)  $\delta$  186.7, 180.4, 170.1, 161.2, 142.5, 137.2, 130.7, 129.6, 128.8, 127.8, 126.0, 125.7, 123.2, 123.1, 121.8, 110.3, 110.2, 103.5, 53.2, 51.7, 51.5, 50.4, 39.6, 20.8.

**IR**: 3363, 2953, 2350, 1728, 1619, 1470, 1342, 1260, 1115, 752 cm<sup>-1</sup>.

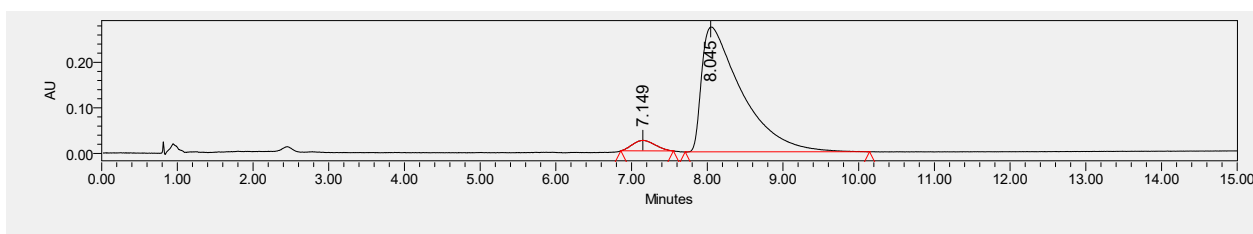
**HRMS** (FTMS+c ESI)  $m/z$ : [M - H]<sup>-</sup> calcd for C<sub>24</sub>H<sub>22</sub>N<sub>2</sub>O<sub>6</sub> 433.1405; found 433.1401.

The UPCC chromatograms of racemic product **C37**

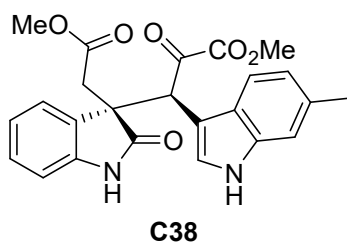


|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 5.634          | 1421522  | 6.00   | 69616  |
| 2 | 6.153          | 1276372  | 5.39   | 47311  |
| 3 | 7.002          | 10469368 | 44.18  | 422858 |
| 4 | 8.006          | 10529067 | 44.43  | 279540 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 7.149          | 474045   | 4.49   | 23247  |
| 2 | 8.045          | 10087822 | 95.51  | 274123 |



**Methyl (S)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-3-(6-methyl-1H-indol-3-yl)-2-oxopropanoate (C38)**

Yellow oil; 39.1 mg, 90% yield, >19:1 dr, 95% ee; [ $\alpha$ ]<sub>D</sub><sup>19.4</sup> = -234.48 ( $c$  = 0.52 in CH<sub>2</sub>Cl<sub>2</sub>).

**UPCC** DAICEL CHIRALCEL OX-3, CO<sub>2</sub>/MeOH = 80/20, flow rate = 1.5 mL/min,  $\lambda$  = 211 nm,  $t_R$  (major) = 9.50 min,  $t_R$  (minor) = 14.90 min.

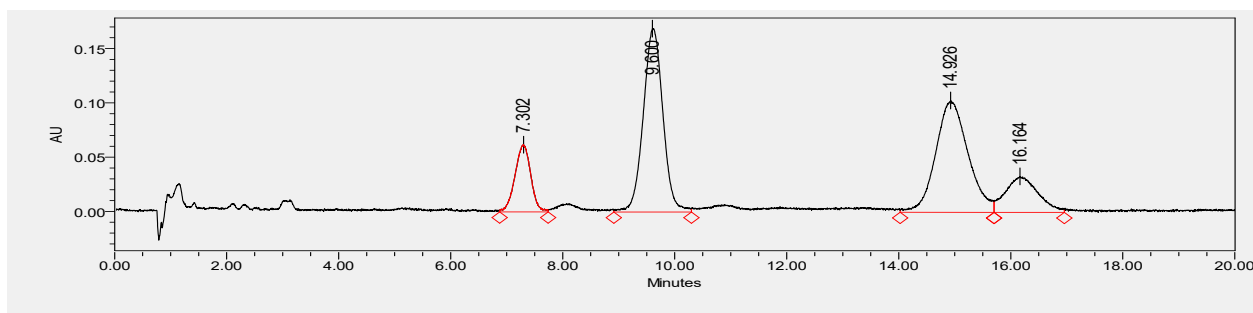
**<sup>1</sup>H NMR** (400 MHz, Chloroform-*d*)  $\delta$  8.62 (s, 1H), 8.49 (s, 1H), 7.54 (d,  $J$  = 8.0 Hz, 1H), 7.23 – 7.13 (m, 2H), 7.01 (dd,  $J$  = 8.4, 1.2 Hz, 1H), 6.92 – 6.82 (m, 2H), 6.77 – 6.69 (m, 2H), 5.46 (s, 1H), 3.55 (s, 3H), 3.33 (s, 3H), 3.16 (dd,  $J$  = 44.0, 16.0 Hz, 2H), 2.44 (s, 3H).

$^{13}\text{C}$  NMR (101 MHz,  $\text{CDCl}_3$ )  $\delta$  187.7, 180.1, 170.0, 161.0, 142.3, 137.0, 132.9, 129.4, 128.7, 126.3, 125.7, 125.0, 122.7, 121.7, 118.7, 111.7, 110.0, 103.7, 53.1, 51.7, 51.5, 50.1, 40.2, 21.8.

IR: 3365, 2953, 2349, 1729, 1620, 1470, 1342, 1263, 1116, 756  $\text{cm}^{-1}$ .

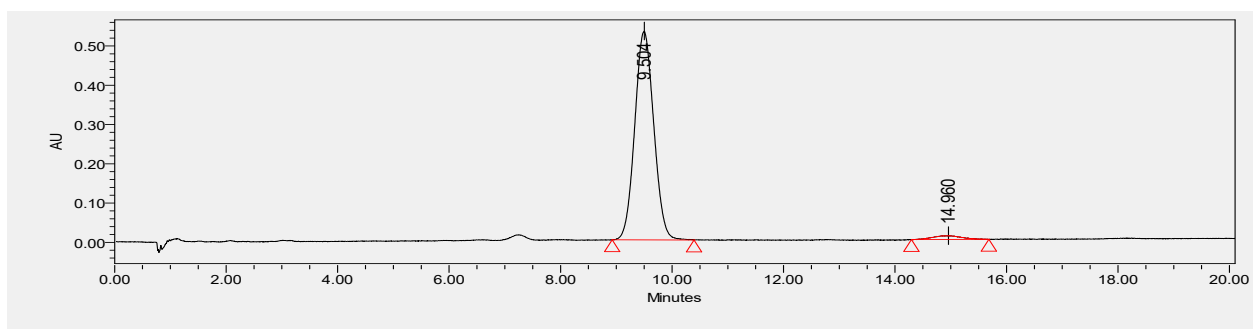
HRMS (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{24}\text{H}_{22}\text{N}_2\text{O}_6$  433.1405; found 433.1401.

The UPCC chromatograms of racemic product **C38**

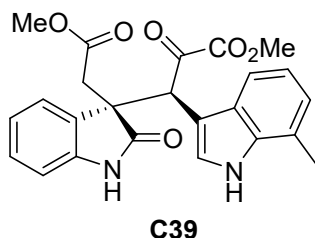


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 7.302          | 1209847 | 11.21  | 62003  |
| 2 | 9.600          | 4126123 | 38.25  | 168907 |
| 3 | 14.926         | 4123306 | 38.22  | 103108 |
| 4 | 16.164         | 1328557 | 12.32  | 33166  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 9.504          | 12569746 | 97.41  | 531700 |
| 2 | 14.960         | 333726   | 2.59   | 9749   |



**Methyl (S)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-3-(7-methyl-1H-indol-3-yl)-2-oxopropanoate (C39)**

Yellow oil; 33.9 mg, 78% yield, >19:1 dr, 96% ee;  $[\alpha]_{\text{D}}^{19.5} = -319.57$  ( $c = 0.14$  in  $\text{CH}_2\text{Cl}_2$ ).

UPCC DAICEL CHIRALCEL IE-3,  $\text{CO}_2/i\text{-PrOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_{\text{R}}$  (major) = 13.39 min,  $t_{\text{R}}$  (minor) = 12.66 min.

$^1\text{H}$  NMR (400 MHz,  $\text{Chloroform-d}$ )  $\delta$  8.60 (s, 1H), 8.36 (s, 1H), 7.53 (d,  $J = 7.6$  Hz, 1H), 7.19 (t,  $J = 8.0$  Hz, 1H), 7.12 (t,  $J = 7.6$  Hz, 1H), 7.04 (d,  $J = 7.2$  Hz, 1H), 6.89 – 6.84 (m, 3H), 6.72 (d,  $J = 7.2$  Hz, 1H), 5.48 (s, 1H), 3.55 (s, 3H), 3.33 (s, 3H), 3.15 (dd,  $J = 56.0, 16.0$  Hz, 2H), 2.46 (s, 3H).

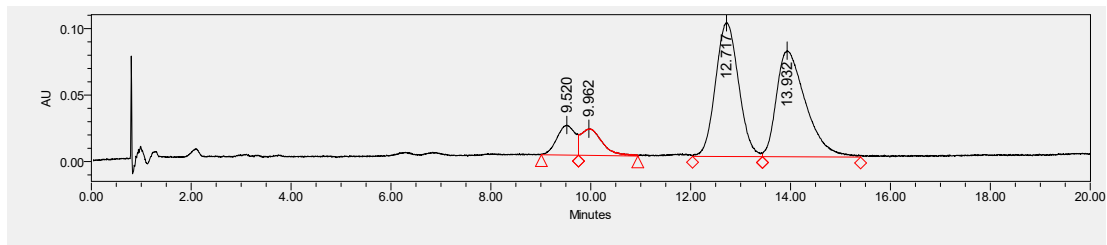
$^{13}\text{C}$  NMR (101 MHz,  $\text{CDCl}_3$ )  $\delta$  187.9, 180.0, 169.9, 160.9, 142.3, 136.2, 129.3, 128.7, 127.4, 126.5, 125.0, 123.6, 121.6, 121.1, 121.0,

116.8, 110.0, 104.5, 53.1, 51.7, 51.5, 50.2, 40.4, 16.6.

IR: 3365, 2953, 2349, 1729, 1620, 1470, 1342, 1263, 1116, 756  $\text{cm}^{-1}$ .

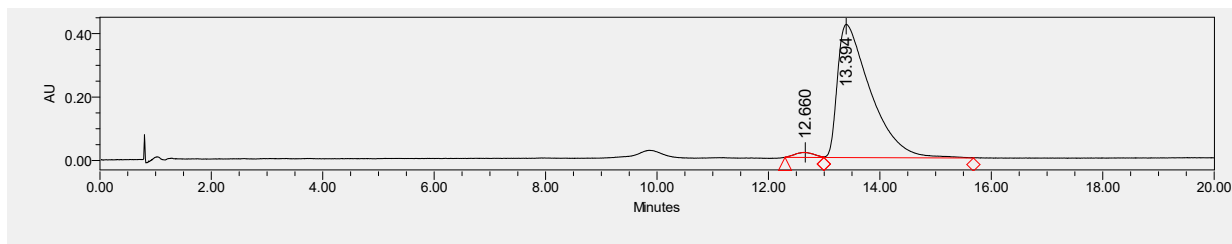
HRMS (FTMS+c ESI)  $m/z$ :  $[M - H]^-$  calcd for  $\text{C}_{24}\text{H}_{22}\text{N}_2\text{O}_6$  433.1405; found 433.1401.

The UPCC chromatograms of racemic product **C39**

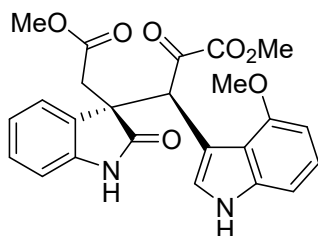


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 9.520          | 569397  | 7.47   | 22627  |
| 2 | 9.962          | 595313  | 7.81   | 20096  |
| 3 | 12.717         | 3224534 | 42.32  | 100929 |
| 4 | 13.932         | 3229761 | 42.39  | 79839  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 12.660         | 381741   | 2.16   | 16001  |
| 2 | 13.394         | 17306741 | 97.84  | 420302 |



**C40**

**Methyl (S)-3-(4-methoxy-1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (C40)**

Yellow oil; 40.1 mg, 91% yield, >19:1 dr, 90% ee;  $[\alpha]_D^{19.2} = -147.39$  ( $c = 0.80$  in  $\text{CH}_2\text{Cl}_2$ ).

**UPCC** DAICEL CHIRALCEL OD-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm,  $t_R$  (major) = 2.89 min,  $t_R$  (minor) = 4.89 min.

**$^1\text{H}$  NMR** (400 MHz,  $\text{Chloroform-}d$ )  $\delta$  8.78 (s, 1H), 8.32 (s, 1H), 7.18 (td,  $J = 7.6, 1.2$  Hz, 1H), 7.13 (t,  $J = 8.0$  Hz, 1H), 6.96 (d,  $J = 8.4$  Hz, 1H), 6.88 (d,  $J = 7.7$  Hz, 1H), 6.83 (t,  $J = 7.6$  Hz, 1H), 6.65 (d,  $J = 7.6$  Hz, 1H), 6.62 – 6.51 (m, 2H), 6.06 (s, 1H), 4.01 (s, 3H), 3.55 (s, 3H), 3.37 (d,  $J = 15.6$  Hz, 1H), 3.27 (s, 3H), 3.02 (d,  $J = 15.6$  Hz, 1H).

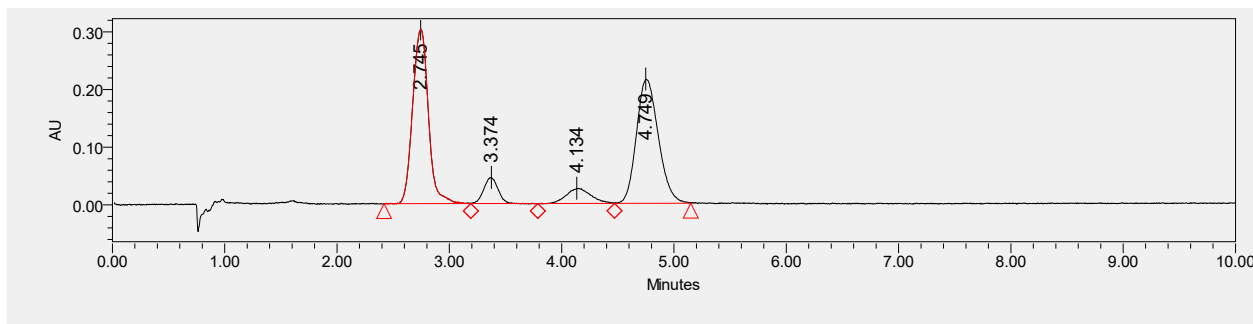
**$^{13}\text{C}$  NMR** (101 MHz,  $\text{CDCl}_3$ )  $\delta$  188.2, 180.5, 170.4, 161.2, 154.6, 142.6, 138.2, 129.5, 128.4, 125.7, 125.2, 123.9, 121.4, 117.6, 109.8,

105.1, 103.8, 100.9, 55.3, 53.0, 51.5, 51.4, 50.8, 40.1.

IR: 3364, 2923, 2350, 1729, 1619, 1468, 1352, 1264, 1090, 756 cm<sup>-1</sup>.

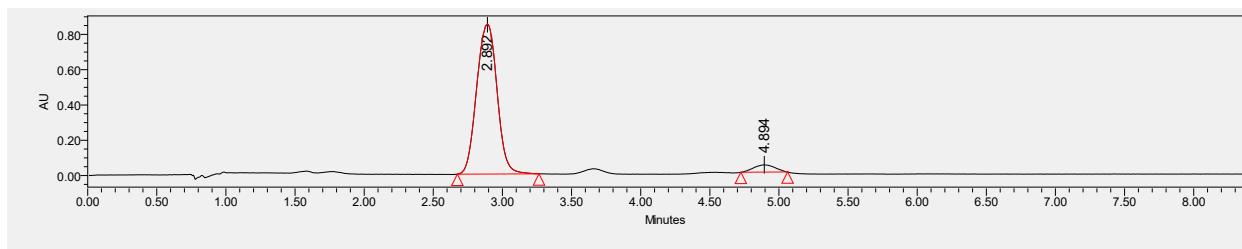
HRMS (FTMS+c ESI) m/z: [M - H]<sup>-</sup> calcd for C<sub>24</sub>H<sub>22</sub>N<sub>2</sub>O<sub>7</sub> 449.1354, found 449.1353.

The UPCC chromatograms of racemic product **C40**

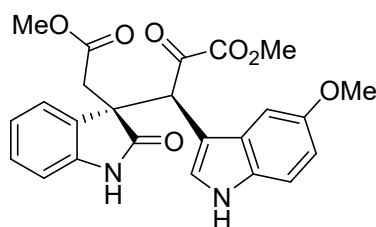


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 2.745          | 2940536 | 44.20  | 303058 |
| 2 | 3.374          | 399988  | 6.01   | 45304  |
| 3 | 4.134          | 395107  | 5.94   | 26191  |
| 4 | 4.749          | 2916474 | 43.84  | 215427 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 2.892          | 8691002 | 95.13  | 851843 |
| 2 | 4.894          | 444798  | 4.87   | 41545  |



**C41**

**Methyl (S)-3-(5-methoxy-1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate (C41)**

Yellow oil; 36.9 mg, 80% yield, >19:1 dr, 95% ee; [ $\alpha$ ]<sub>D</sub><sup>19.5</sup> = -273.39 (c = 0.50 in CH<sub>2</sub>Cl<sub>2</sub>).

UPCC DAICEL CHIRALCEL OX-3, CO<sub>2</sub>/MeOH = 80/20, flow rate = 1.5 mL/min,  $\lambda$  = 211 nm, t<sub>R</sub> (major) = 8.90 min, t<sub>R</sub> (minor) = 11.19 min.

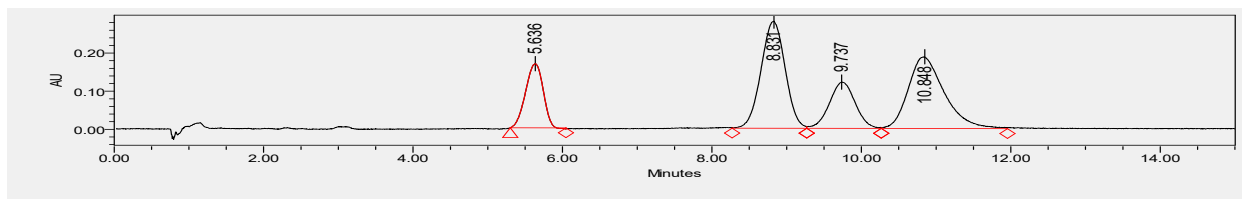
<sup>1</sup>H NMR (400 MHz, Chloroform-*d*)  $\delta$  8.64 (d, *J* = 2.8 Hz, 1H), 8.36 (s, 1H), 7.24 (d, *J* = 8.8 Hz, 1H), 7.19 (td, *J* = 7.6, 1.2 Hz, 1H), 7.04 (d, *J* = 2.4 Hz, 1H), 6.91 – 6.83 (m, 3H), 6.80 (d, *J* = 2.8 Hz, 1H), 6.73 (d, *J* = 7.2 Hz, 1H), 3.83 (s, 3H), 3.56 (s, 3H), 3.35 (s, 3H), 3.12 (dd, *J* = 56.0, 12.0 Hz, 2H).

<sup>13</sup>C NMR (101 MHz, CDCl<sub>3</sub>)  $\delta$  187.8, 180.0, 169.9, 161.0, 155.0, 142.3, 131.6, 129.4, 128.7, 128.3, 127.4, 125.0, 121.7, 113.5, 112.6, 110.0, 103.6, 100.4, 56.0, 53.2, 51.8, 51.6, 50.1, 40.4.

IR: 3365, 2953, 2350, 1731, 1621, 1487, 1352, 1216, 1164, 752  $\text{cm}^{-1}$ .

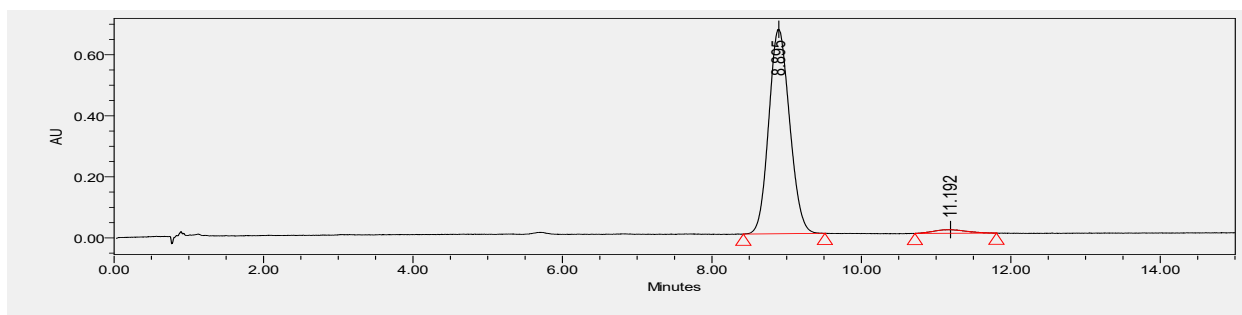
HRMS (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{24}\text{H}_{22}\text{N}_2\text{O}_7$  449.1354, found 449.1353.

The UPCC chromatograms of racemic product **C41**

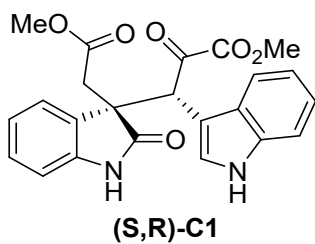


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 5.636          | 2763922 | 15.15  | 168553 |
| 2 | 8.831          | 6283662 | 34.43  | 281068 |
| 3 | 9.737          | 2910284 | 15.95  | 121528 |
| 4 | 10.848         | 6290603 | 34.47  | 188066 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 8.895          | 12972797 | 97.32  | 669967 |
| 2 | 11.192         | 357510   | 2.68   | 12049  |

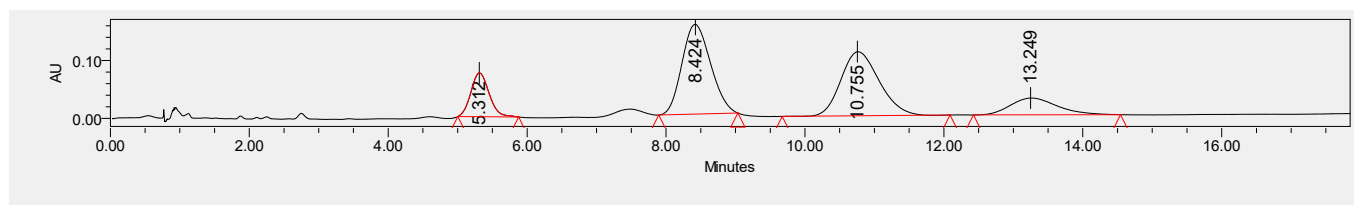


**Methyl (R)-3-(1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-2-oxopropanoate ((S,R)-C1)**

Yellow oil; 40.0 mg, 95% yield, 85:15 dr, 89%/92% ee;  $[\alpha]_D^{21.9} = 114.08$  ( $c = 0.28$  in  $\text{CH}_2\text{Cl}_2$ )

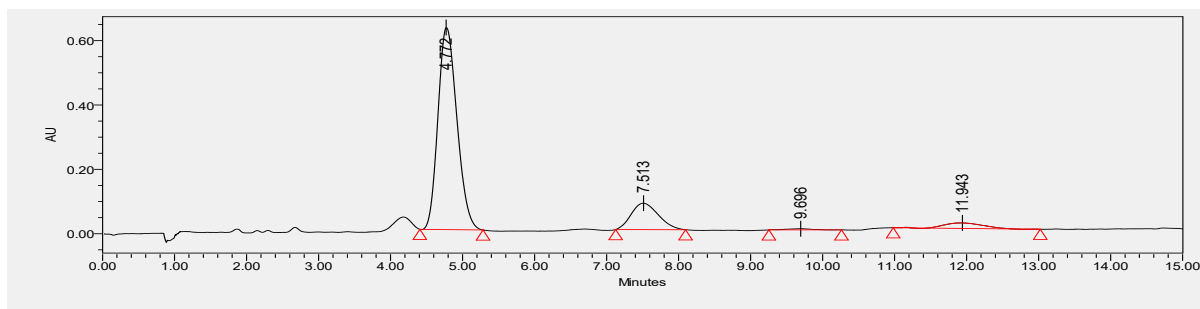
**UPCC** DAICEL CHIRALCEL OZ-3,  $\text{CO}_2/\text{MeOH} = 80/20$ , flow rate = 1.5 mL/min,  $\lambda = 211$  nm, major isomer:  $t_R$  (major) = 4.77 min,  $t_R$  (minor) = 11.94 min; minor isomer :  $t_R$  (major) = 7.51 min,  $t_R$  (minor) = 9.70 min.

The UPCC chromatograms of racemic product **C1**

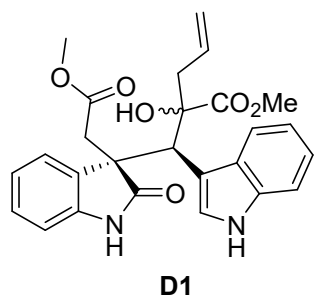


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 5.312          | 1379872 | 12.09  | 75677  |
| 2 | 8.424          | 4442531 | 38.92  | 155097 |
| 3 | 10.755         | 4234268 | 37.10  | 110819 |
| 4 | 13.249         | 1356487 | 11.89  | 29024  |

The UPCC chromatograms of **(S,R)-C1**



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 4.772          | 11493953 | 79.86  | 627867 |
| 2 | 7.513          | 2145997  | 14.91  | 82073  |
| 3 | 9.696          | 85913    | 0.60   | 3690   |
| 4 | 11.943         | 667511   | 4.64   | 17122  |



**Methyl 2-((S)-3-(1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)methyl)-2-hydroxypent-4-enoate (D1)**

Yellow solid; 36.9 mg, 87% yield, 94:6 dr, 95%/99% ee.

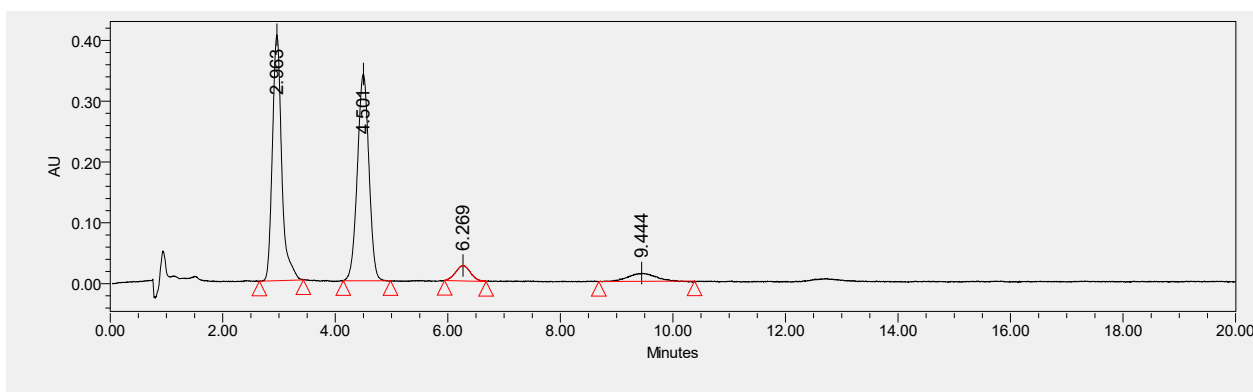
**UPCC** DAICEL CHIRALCEL IH-3, CO<sub>2</sub>/MeOH = 80/20, flow rate = 1.5 mL/min,  $\lambda$  = 211 nm, major isomer:  $t_R$  (major) = 2.96 min,  $t_R$  (minor) = 4.48 min; minor isomer:  $t_R$  (major) = 6.25.

<sup>1</sup>H NMR (400 MHz, DMSO-*d*<sub>6</sub>)  $\delta$  10.91 (s, 1H), 10.79 (s, 1H), 7.40 (d,  $J$  = 7.8 Hz, 1H), 7.35 (d,  $J$  = 7.4 Hz, 1H), 7.23 – 7.14 (m, 2H), 6.96 – 6.80 (m, 3H), 6.70 (t,  $J$  = 7.6 Hz, 1H), 6.45 (d,  $J$  = 7.6 Hz, 1H), 6.08 – 6.01 (m, 1H), 5.55 – 5.46 (m, 1H), 4.83 (d,  $J$  = 11.6 Hz, 1H), 4.72 (d,  $J$  = 17.2 Hz, 1H), 4.17 (s, 1H), 3.69 (s, 3H), 3.20 (s, 3H), 3.05 – 2.88 (m, 2H), 1.80 (ddd,  $J$  = 76.4, 13.6, 7.6 Hz, 2H).

<sup>13</sup>C NMR (101 MHz, DMSO)  $\delta$  182.1, 176.1, 169.1, 141.7, 134.4, 132.8, 131.6, 128.7, 127.8, 123.4, 121.3, 120.6, 118.5, 117.8, 117.6, 111.1, 109.0, 108.6, 80.7, 55.1, 51.9, 51.3, 46.7, 44.7.

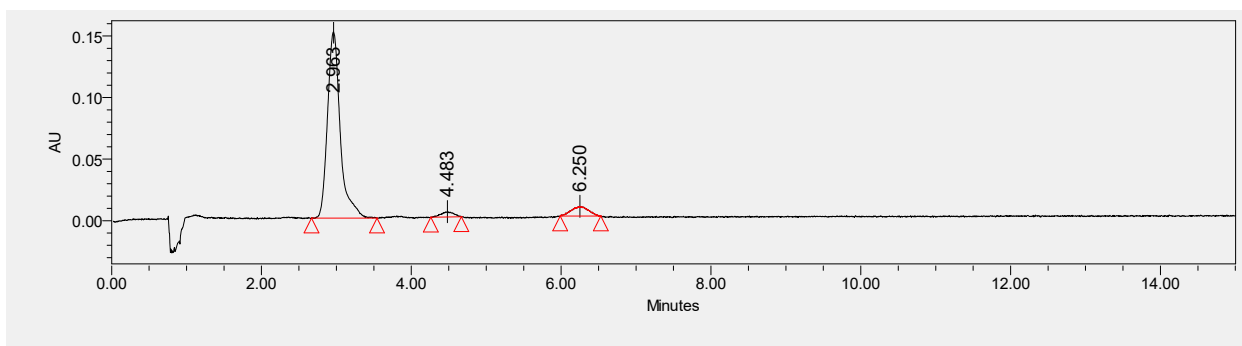
**HRMS** (FTMS+c ESI)  $m/z$ : [M - H]<sup>-</sup> calcd for C<sub>26</sub>H<sub>26</sub>N<sub>2</sub>O<sub>6</sub> 461.1718; found 461.1720.

The UPCC chromatograms of racemic product **D1**

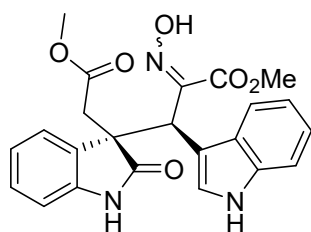


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 2.963          | 4632491 | 44.52  | 404326 |
| 2 | 4.501          | 4841372 | 46.53  | 339975 |
| 3 | 6.269          | 467365  | 4.49   | 25437  |
| 4 | 9.444          | 464386  | 4.46   | 13607  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 2.963          | 1787069 | 91.12  | 151218 |
| 2 | 4.483          | 49827   | 2.54   | 4328   |
| 3 | 6.250          | 124305  | 6.34   | 7858   |



**D2**

**Methyl (S)-2-(hydroxyimino)-3-(1H-indol-3-yl)-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)propanoate (D2)**

Yellow oil; 21.8 mg, 50% yield, >19:1 dr, >19/1 Z/E, 95% ee.

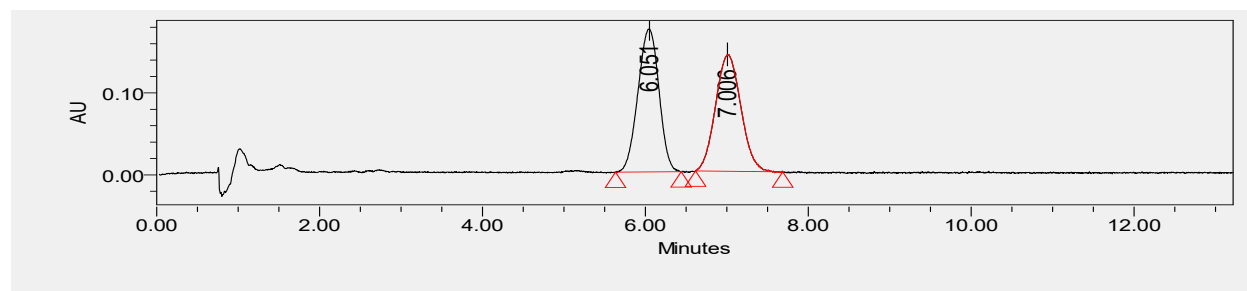
**UPCC** DAICEL CHIRALCEL IH-3, CO<sub>2</sub>/MeOH = 80/20, flow rate = 1.5 mL/min,  $\lambda$  = 211 nm,  $t_R$  (major) = 6.98 min,  $t_R$  (minor) = 6.03 min.

<sup>1</sup>H NMR (400 MHz, Chloroform-*d*)  $\delta$  11.13 (s, 1H), 8.61 (s, 1H), 8.47 (s, 1H), 7.64 (d,  $J$  = 8.0 Hz, 1H), 7.36 (d,  $J$  = 8.0 Hz, 1H), 7.23 – 7.06 (m, 4H), 6.92 – 6.77 (m, 3H), 4.96 (s, 1H), 3.59 (s, 3H), 3.31 – 3.27 (m, 4H), 2.97 (d,  $J$  = 16.0 Hz, 1H).

<sup>13</sup>C NMR (101 MHz, CDCl<sub>3</sub>)  $\delta$  180.8, 170.1, 163.2, 148.2, 142.3, 135.8, 129.9, 128.5, 128.0, 126.5, 124.7, 122.4, 121.7, 120.3, 119.0, 111.6, 110.1, 108.8, 53.5, 52.7, 51.7, 43.3, 41.5.

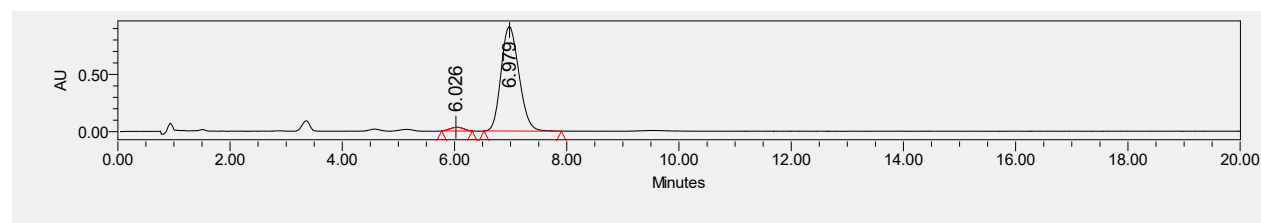
**HRMS** (FTMS+c ESI)  $m/z$ : [M + Na]<sup>+</sup> calcd for C<sub>23</sub>H<sub>21</sub>N<sub>3</sub>O<sub>6</sub> 458.1323; found 458.1327.

The UPCC chromatograms of racemic product **D2**

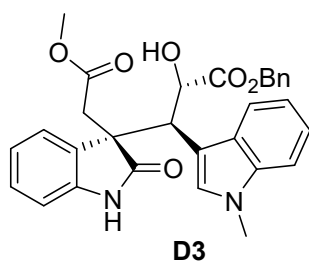


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 6.051          | 3127827 | 50.48  | 174597 |
| 2 | 7.006          | 3068427 | 49.52  | 142419 |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area     | % Area | Height |
|---|----------------|----------|--------|--------|
| 1 | 6.026          | 544255   | 2.62   | 32628  |
| 2 | 6.979          | 20199744 | 97.38  | 912684 |



**Benzyl (2S,3S)-2-hydroxy-3-((S)-3-(2-methoxy-2-oxoethyl)-2-oxoindolin-3-yl)-3-(1-methyl-1H-indol-3-yl)propanoate (D3)**

Yellow oil; 37.4 mg, 73% yield, >19:1 dr, 98% ee.

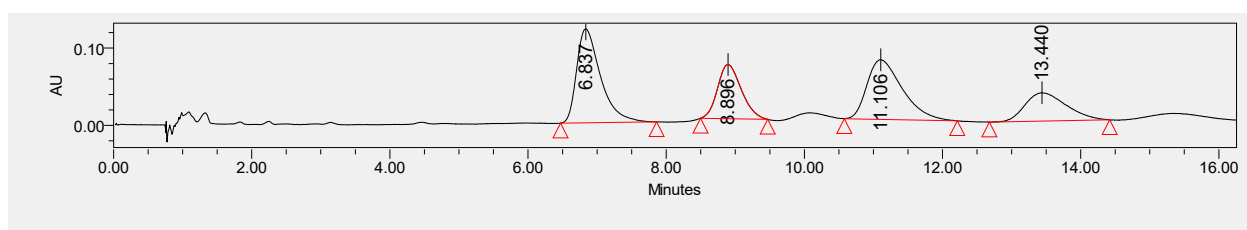
**UPCC** DAICEL CHIRALCEL IA-3, CO<sub>2</sub>/MeOH = 80/20, flow rate = 1.5 mL/min, λ = 211 nm, major isomer: t<sub>R</sub> (major) = 6.61 min, t<sub>R</sub> (minor) = 11.18 min; minor isomer: t<sub>R</sub> (major) = 8.88 min, t<sub>R</sub> (minor) = 13.41 min.

<sup>1</sup>H NMR (400 MHz, Chloroform-*d*) δ 8.56 (s, 1H), 7.35 (d, *J* = 7.9 Hz, 1H), 7.27 – 7.18 (m, 3H), 7.15 – 7.06 (m, 5H), 7.00 – 6.91 (m, 2H), 6.90 – 6.84 (m, 2H), 6.70 (d, *J* = 7.8 Hz, 1H), 4.88 (t, *J* = 2.4 Hz, 1H), 4.82 (d, *J* = 12.0 Hz, 1H), 4.60 (d, *J* = 12.0 Hz, 1H), 4.33 (d, *J* = 2.4 Hz, 1H), 4.18 (d, *J* = 2.4 Hz, 1H), 3.65 (s, 3H), 3.33 (s, 3H), 3.16 (dd, *J* = 77.6, 16.0 Hz, 2H).

<sup>13</sup>C NMR (101 MHz, CDCl<sub>3</sub>) δ 182.0, 172.9, 170.2, 141.5, 135.8, 134.9, 131.0, 129.4, 129.0, 128.5, 128.4, 128.4, 128.2, 124.1, 122.0, 121.4, 119.3, 118.7, 109.9, 109.1, 106.3, 72.0, 67.5, 54.3, 51.7, 44.6, 40.6, 33.0.

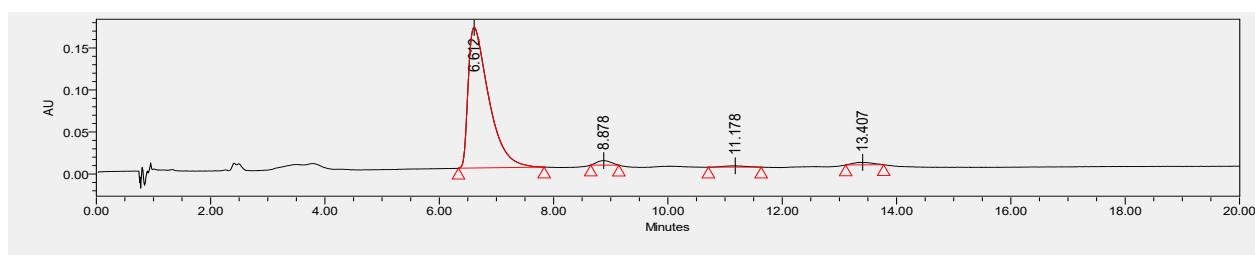
**HRMS** (FTMS+c ESI) *m/z*: [M - H]<sup>-</sup> calcd for C<sub>30</sub>H<sub>28</sub>N<sub>2</sub>O<sub>6</sub> 511.1875; found 511.1877.

The UPCC chromatograms of racemic product **D3**

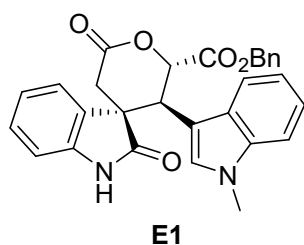


|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 6.837          | 2920994 | 33.01  | 121506 |
| 2 | 8.896          | 1672674 | 18.90  | 70130  |
| 3 | 11.106         | 2735874 | 30.92  | 77152  |
| 4 | 13.440         | 1518685 | 17.16  | 36676  |

The UPCC chromatograms of product from asymmetric reaction



|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 6.612          | 4173564 | 95.42  | 167388 |
| 2 | 8.878          | 89029   | 2.04   | 5374   |
| 3 | 11.178         | 41839   | 0.96   | 1655   |
| 4 | 13.407         | 69348   | 1.59   | 2931   |



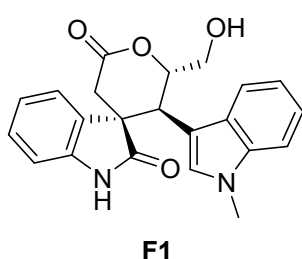
**Benzyl (2'S,3S,3'S)-3'-(1-methyl-1H-indol-3-yl)-2,6'-dioxo-2',3',5',6'-tetrahydrospiro[indoline-3,4'-pyran]-2'-carboxylate (E1)**

Yellow solid; 37.9 mg, 79% yield, >19:1 dr.

$^1\text{H}$  NMR (400 MHz, Chloroform-*d*)  $\delta$  8.15 (s, 1H), 7.35 – 7.29 (m, 2H), 7.22 – 7.16 (m, 1H), 7.14 – 7.05 (m, 4H), 7.03 – 6.92 (m, 3H), 6.86 (s, 1H), 6.69 (d,  $J$  = 7.2 Hz, 2H), 6.59 (d,  $J$  = 8.0 Hz, 1H), 5.95 (d,  $J$  = 11.2 Hz, 1H), 4.76 (s, 2H), 4.19 (d,  $J$  = 11.2 Hz, 1H), 3.35 (s, 3H), 2.99 (dd,  $J$  = 68.0, 20.0 Hz, 2H).

$^{13}\text{C}$  NMR (101 MHz,  $\text{CDCl}_3$ )  $\delta$  179.8, 168.6, 167.6, 140.1, 136.2, 134.6, 130.5, 129.1, 128.4, 128.3, 128.1, 127.6, 126.9, 123.4, 123.0, 121.9, 119.5, 118.4, 110.2, 109.2, 105.2, 78.5, 67.5, 51.2, 39.9, 38.1, 32.7.

HRMS (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{29}\text{H}_{24}\text{N}_2\text{O}_5$  479.1612; found 479.1616.



**(2'S,3S,3'S)-2'-(hydroxymethyl)-3'-(1-methyl-1H-indol-3-yl)-2',3'-dihydrospiro[indoline-3,4'-pyran]-2,6'(5'H)-dione (F1)**

White solid; 31.9 mg, 85% yield, >19:1 dr, >99% ee.

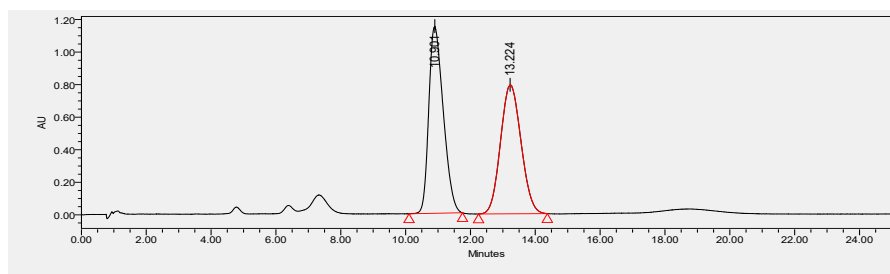
UPCC DAICEL CHIRALCEL IH-3,  $\text{CO}_2/\text{MeOH}$  = 80/20, flow rate = 1.5 mL/min,  $\lambda$  = 211 nm,  $t_R$  (major) = 6.61 min,  $t_R$  (minor) = 11.18 min.

$^1\text{H}$  NMR (400 MHz, Chloroform-*d*)  $\delta$  8.39 (s, 1H), 7.48 (d,  $J$  = 8.0 Hz, 1H), 7.33 (d,  $J$  = 6.8 Hz, 1H), 7.14 – 7.06 (m, 2H), 7.02 – 6.90 (m, 3H), 6.84 (s, 1H), 6.57 (dd,  $J$  = 6.8, 1.6 Hz, 1H), 5.49 (d,  $J$  = 11.2 Hz, 1H), 4.06 (d,  $J$  = 11.3 Hz, 1H), 3.76 (d,  $J$  = 11.2 Hz, 1H), 3.40 (m, 3H), 2.95 (dd,  $J$  = 52.0, 16.0 Hz, 2H), 2.69 (s, 1H).

$^{13}\text{C}$  NMR (101 MHz,  $\text{CDCl}_3$ )  $\delta$  180.5, 169.8, 140.1, 136.3, 131.1, 128.8, 127.7, 126.6, 123.2, 122.9, 121.9, 119.4, 118.6, 110.2, 109.3, 107.5, 81.4, 62.6, 51.3, 38.4, 32.6.

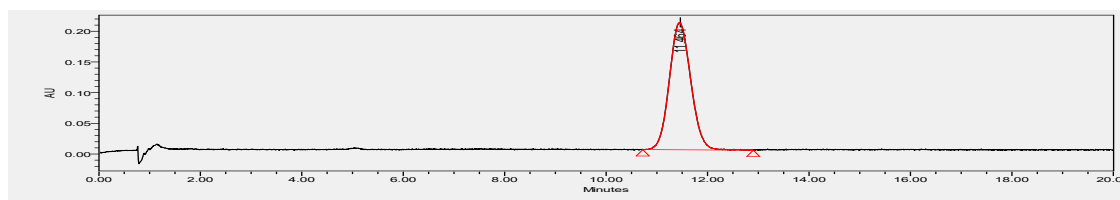
HRMS (FTMS+c ESI)  $m/z$ :  $[\text{M} - \text{H}]^-$  calcd for  $\text{C}_{22}\text{H}_{20}\text{N}_2\text{O}_4$  375.1350; found 375.1351.

The UPCC chromatograms of racemic product **F1**



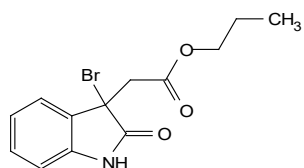
|   | Retention Time | Area     | % Area | Height  |
|---|----------------|----------|--------|---------|
| 1 | 10.901         | 35549445 | 50.56  | 1150128 |
| 2 | 13.224         | 34755554 | 49.44  | 792611  |

The UPCC chromatograms of product from asymmetric reaction



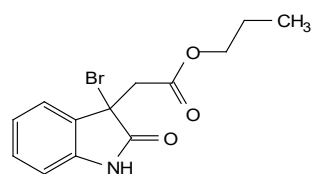
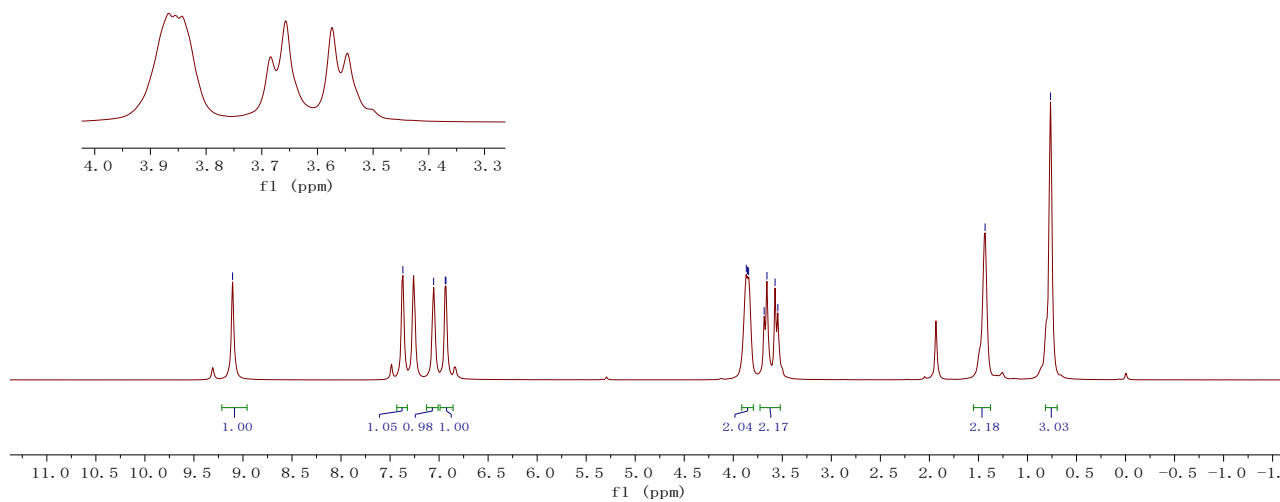
|   | Retention Time | Area    | % Area | Height |
|---|----------------|---------|--------|--------|
| 1 | 11.464         | 6021957 | 100.00 | 207437 |

11 Copies of NMR spectra for products



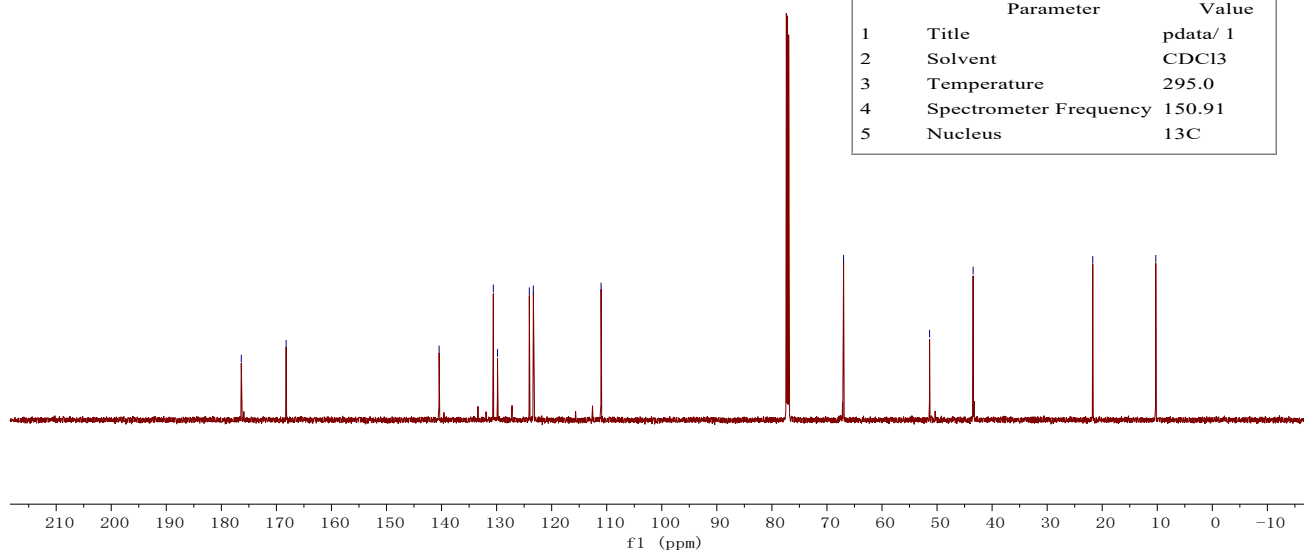
A3

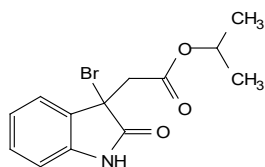
|   | Parameter              | Value          |
|---|------------------------|----------------|
| 1 | Title                  | pdata/ 1       |
| 2 | Solvent                | CDCl3          |
| 3 | Temperature            | 295.0          |
| 4 | Spectrometer Frequency | 600.17         |
| 5 | Nucleus                | <sup>1</sup> H |



A3

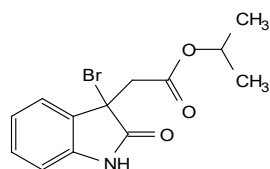
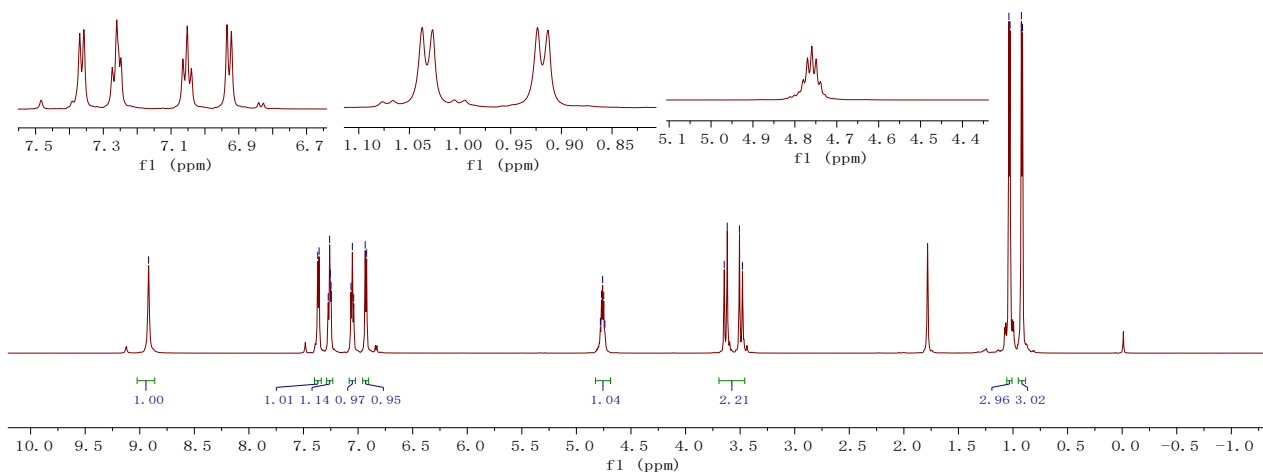
|   | Parameter              | Value           |
|---|------------------------|-----------------|
| 1 | Title                  | pdata/ 1        |
| 2 | Solvent                | CDCl3           |
| 3 | Temperature            | 295.0           |
| 4 | Spectrometer Frequency | 150.91          |
| 5 | Nucleus                | <sup>13</sup> C |





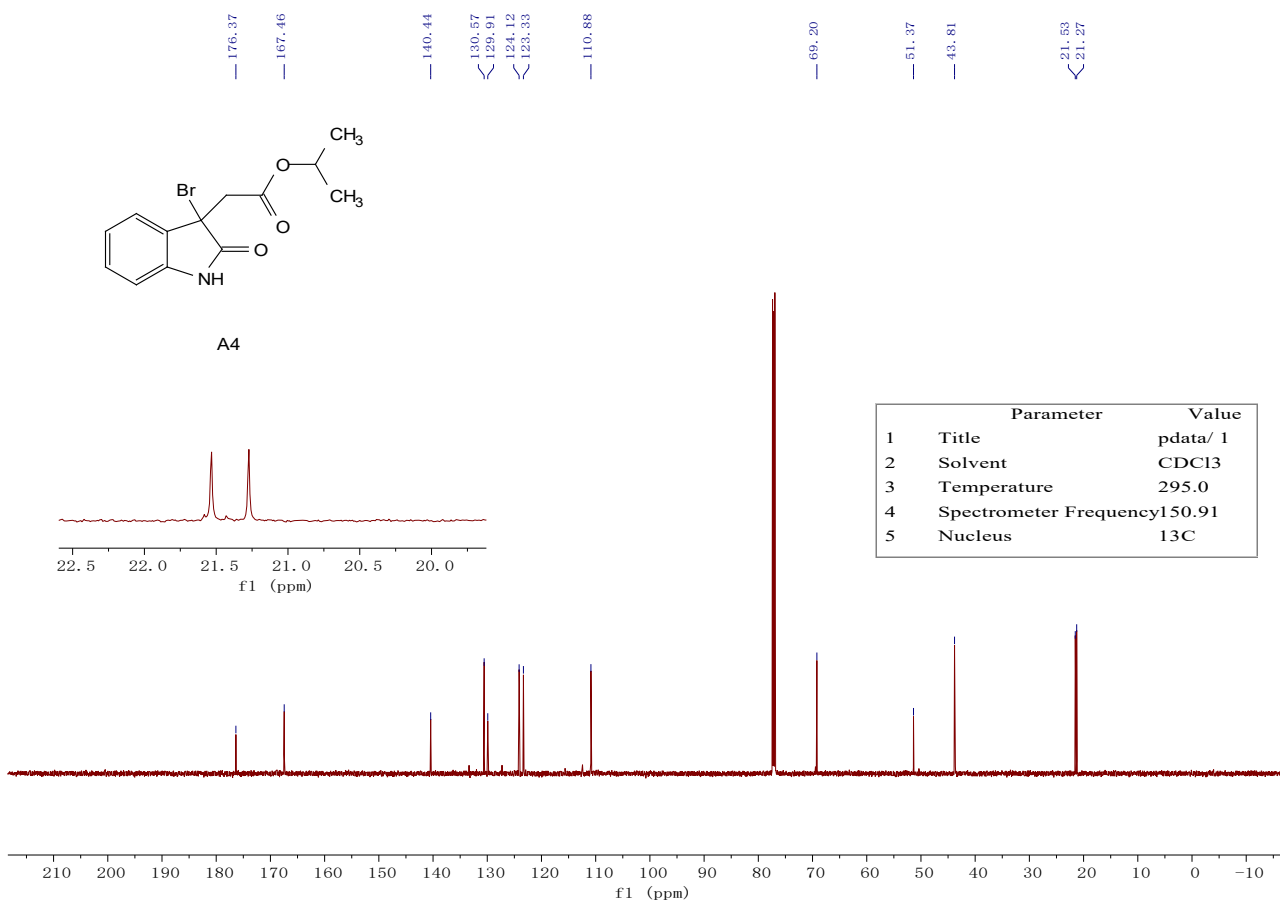
A4

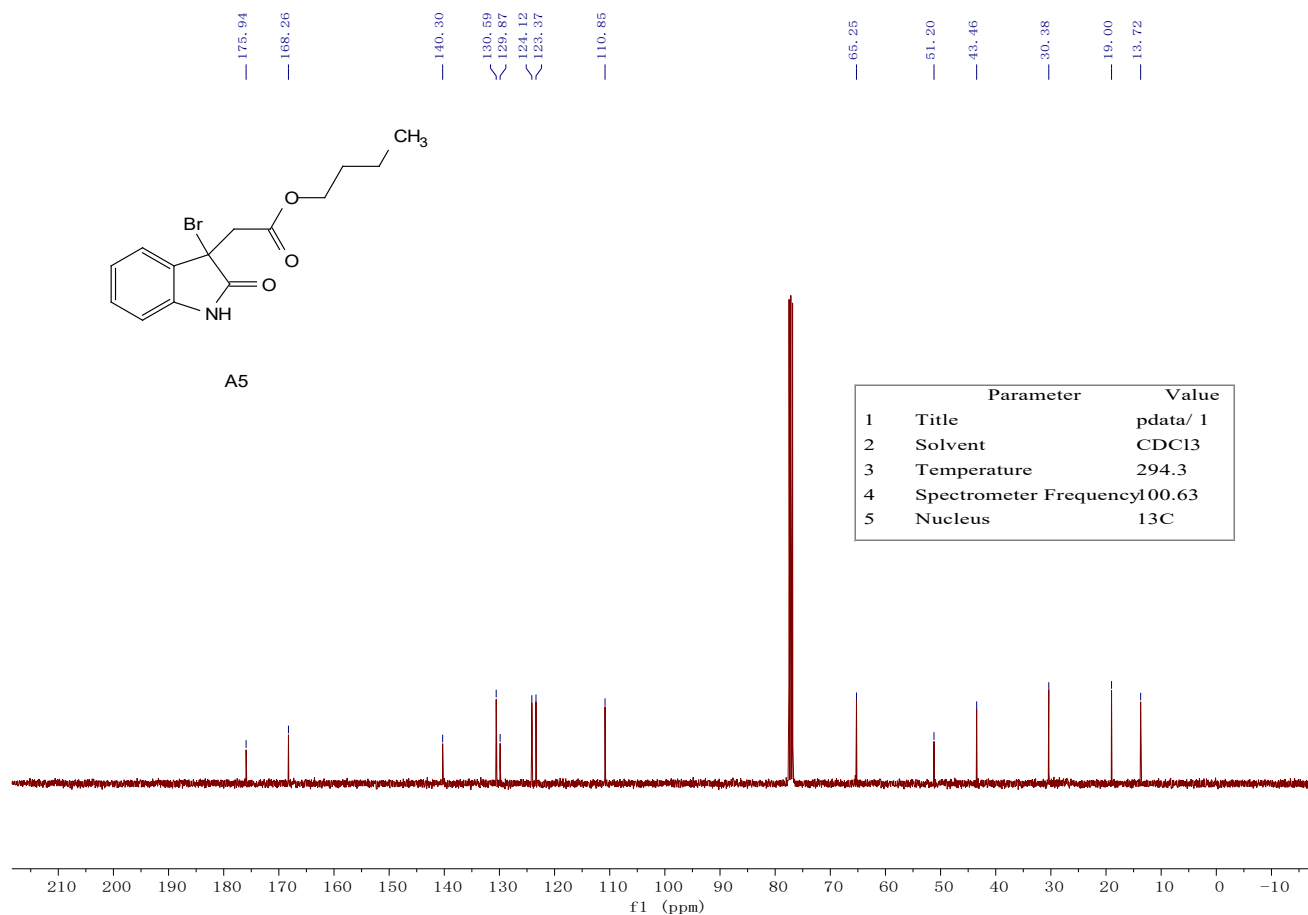
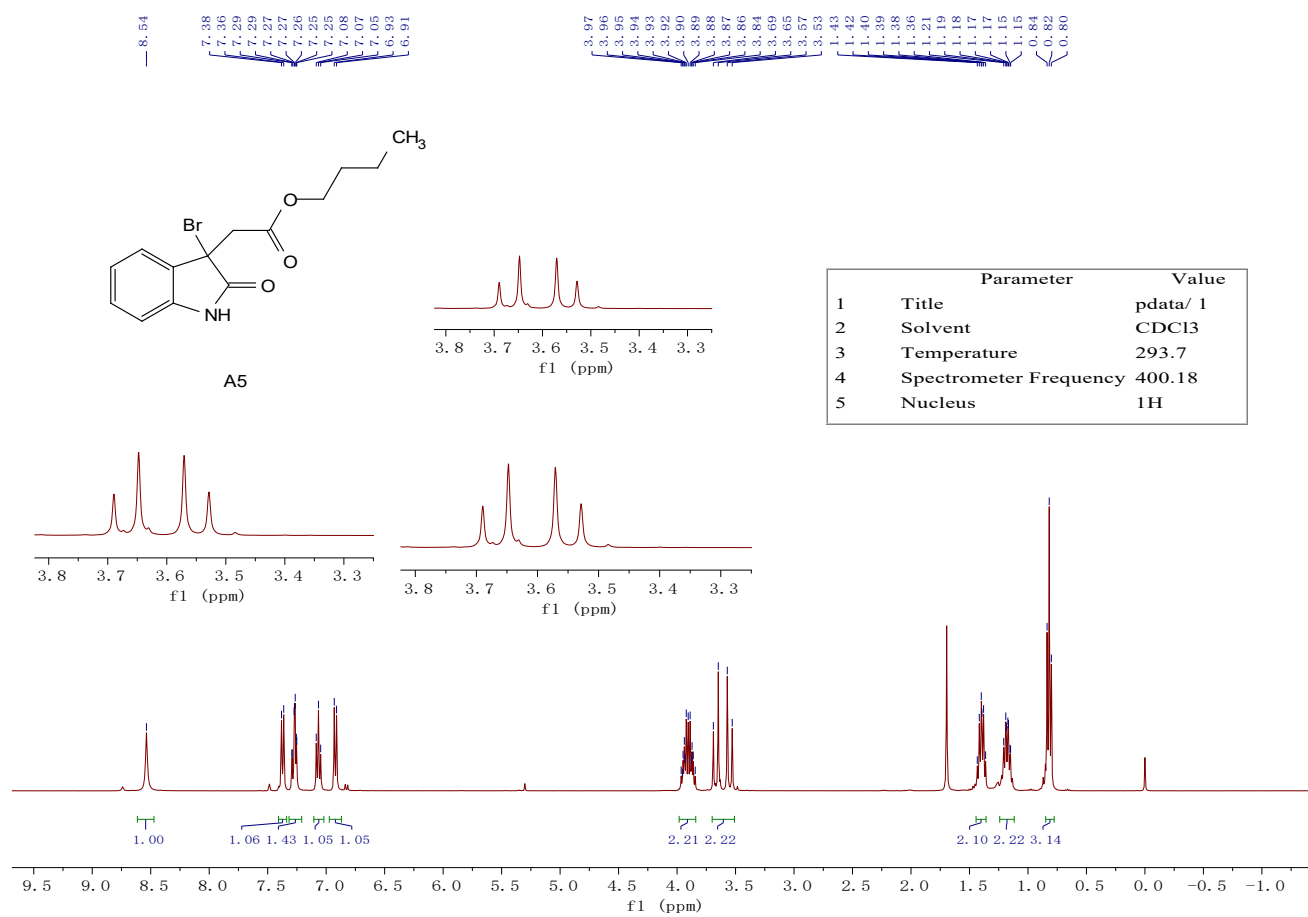
| Parameter                | Value    |
|--------------------------|----------|
| 1 Title                  | pdata/ 1 |
| 2 Solvent                | CDCl3    |
| 3 Temperature            | 295.0    |
| 4 Spectrometer Frequency | 600.17   |
| 5 Nucleus                | 1H       |

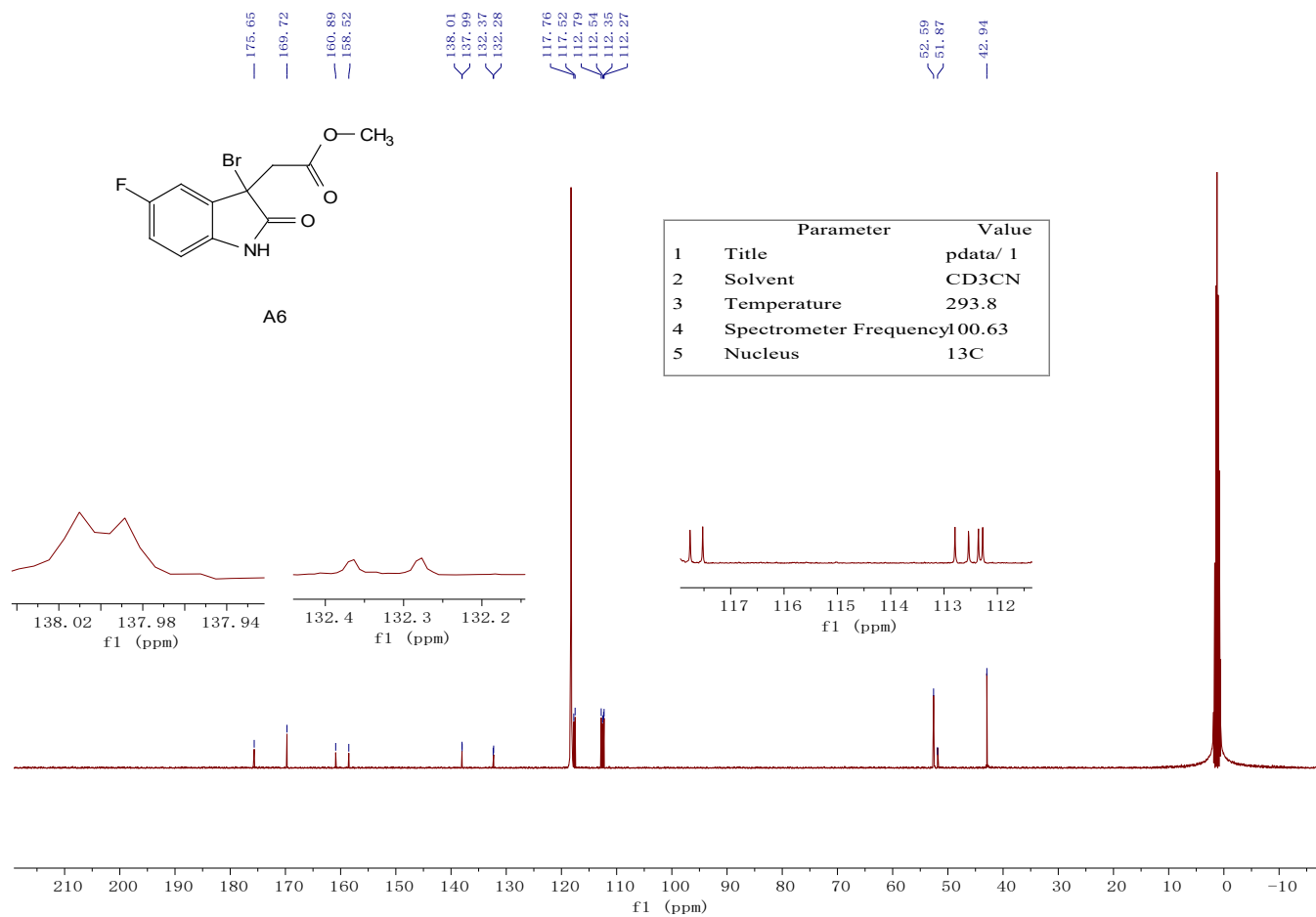
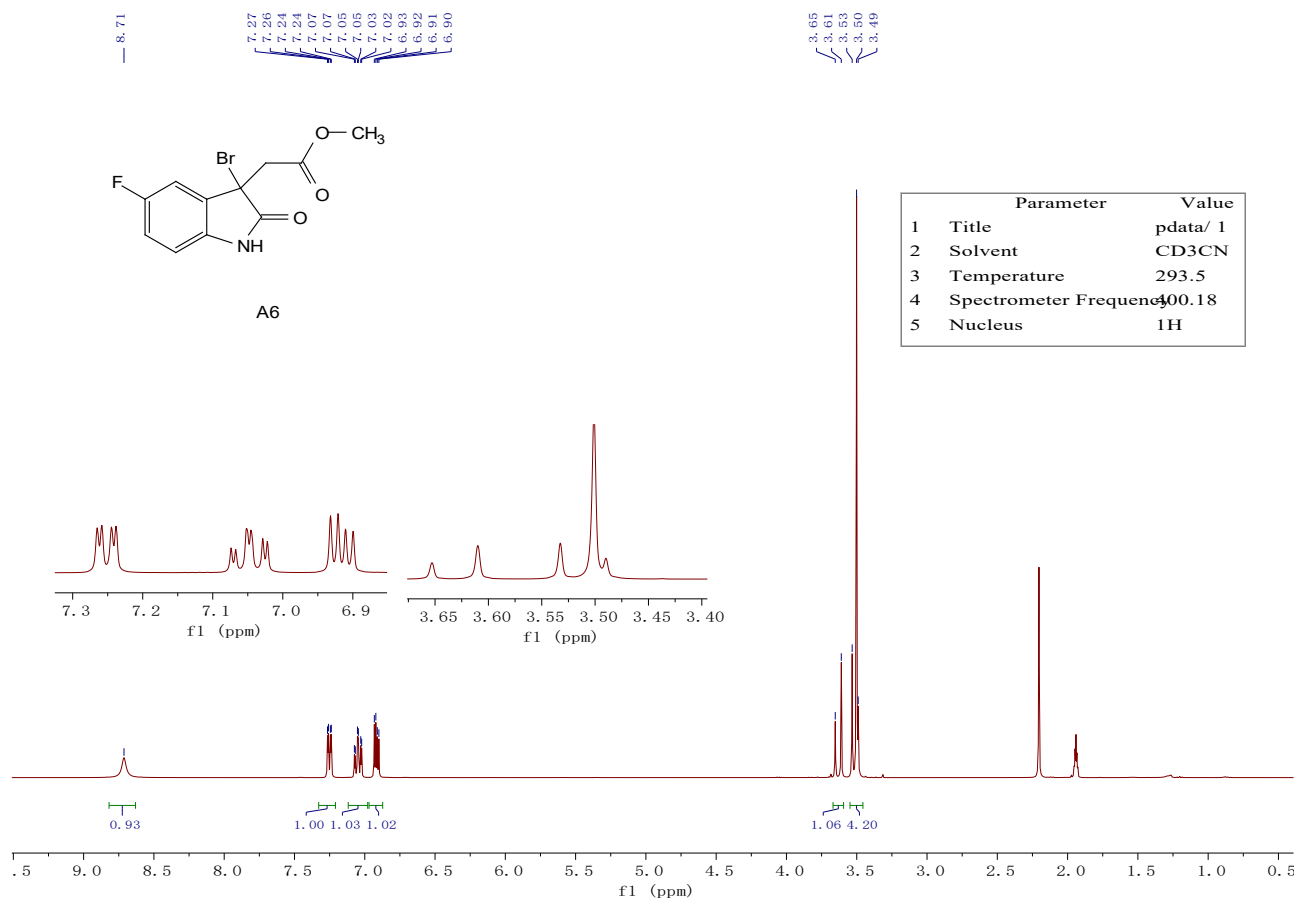


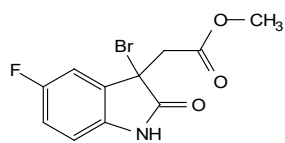
A4

| Parameter                | Value    |
|--------------------------|----------|
| 1 Title                  | pdata/ 1 |
| 2 Solvent                | CDCl3    |
| 3 Temperature            | 295.0    |
| 4 Spectrometer Frequency | 150.91   |
| 5 Nucleus                | 13C      |

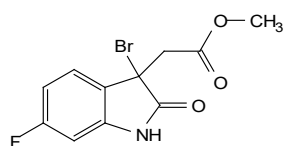
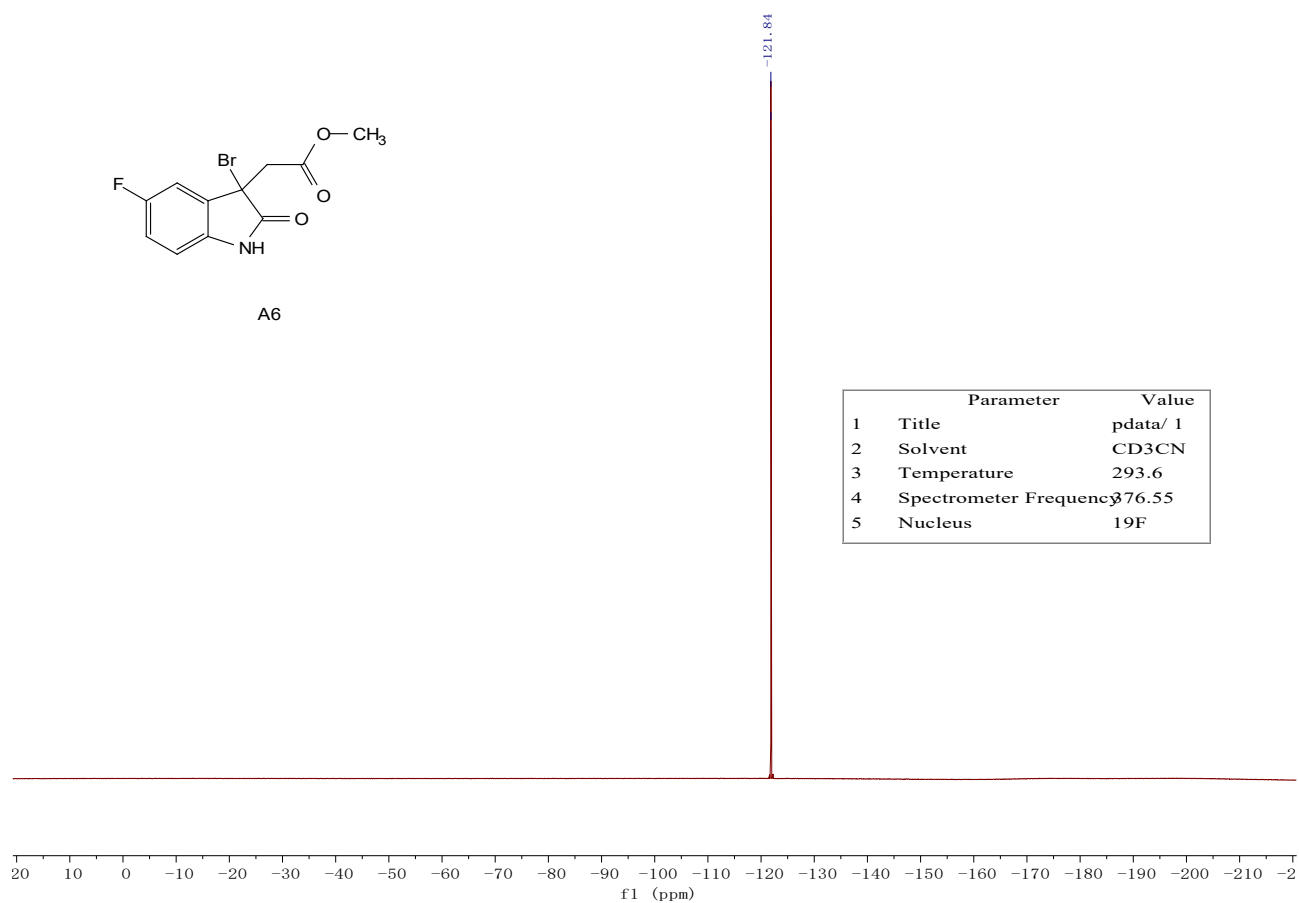




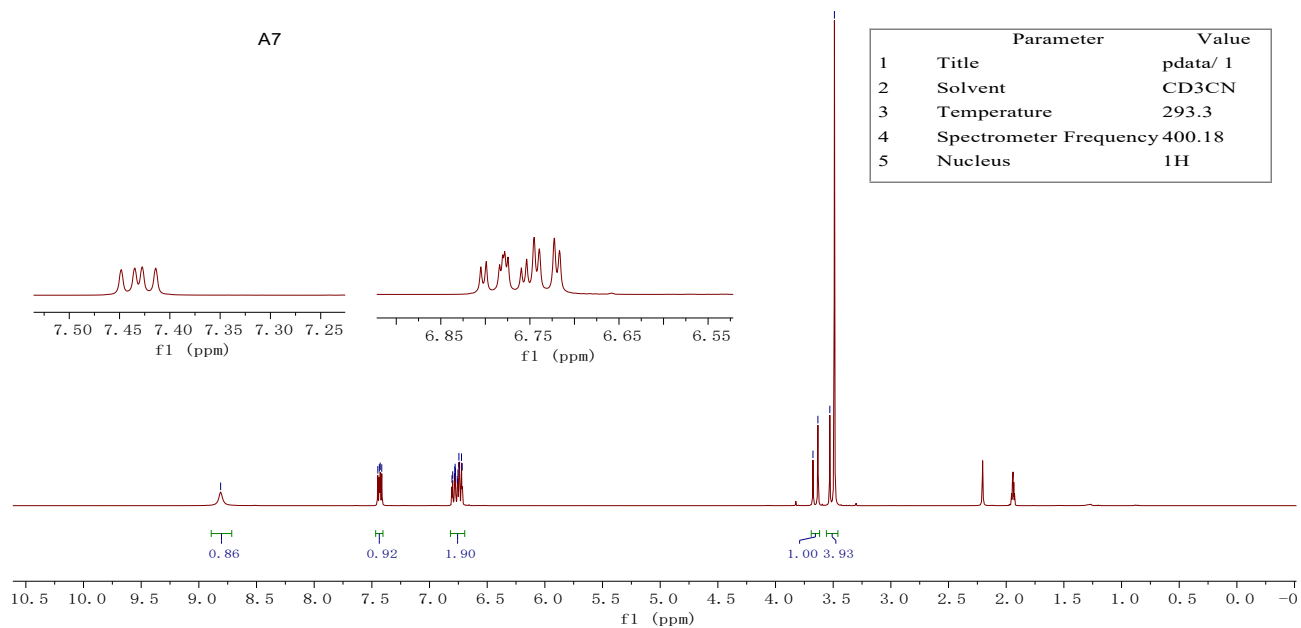


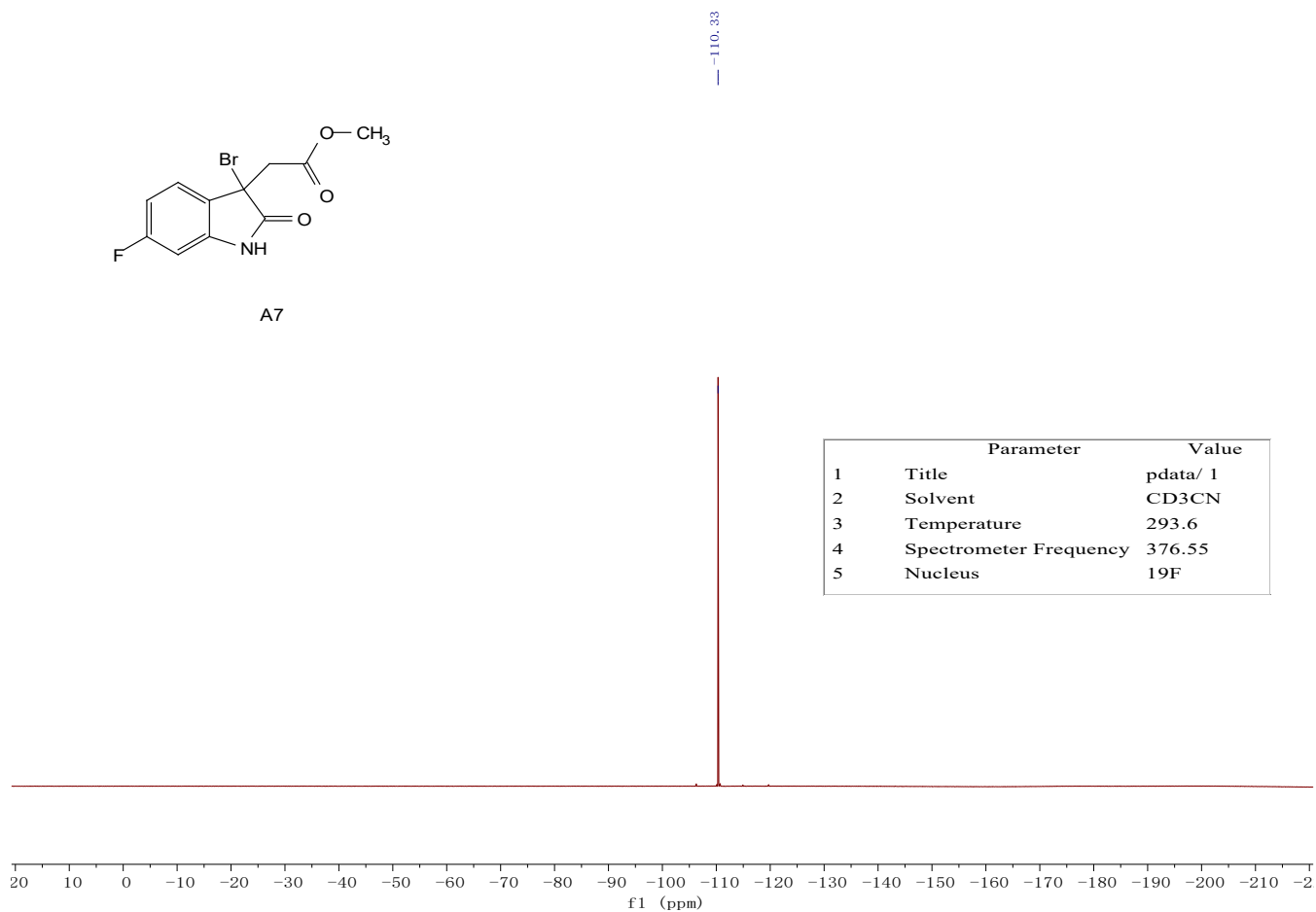
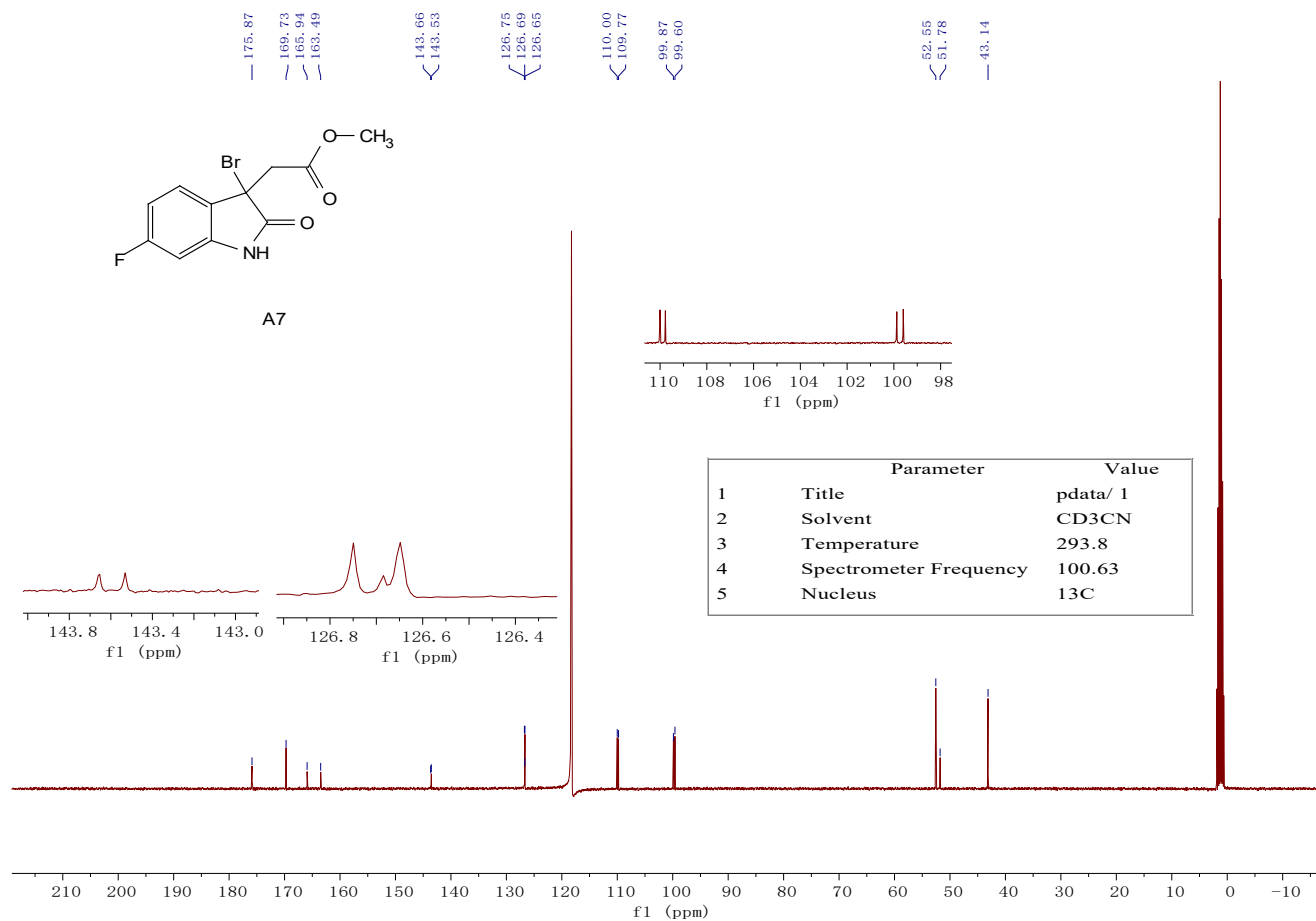


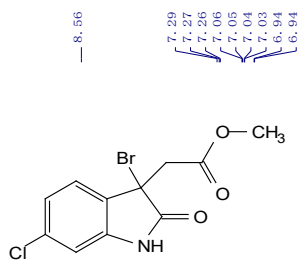
A6



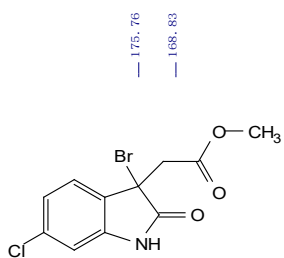
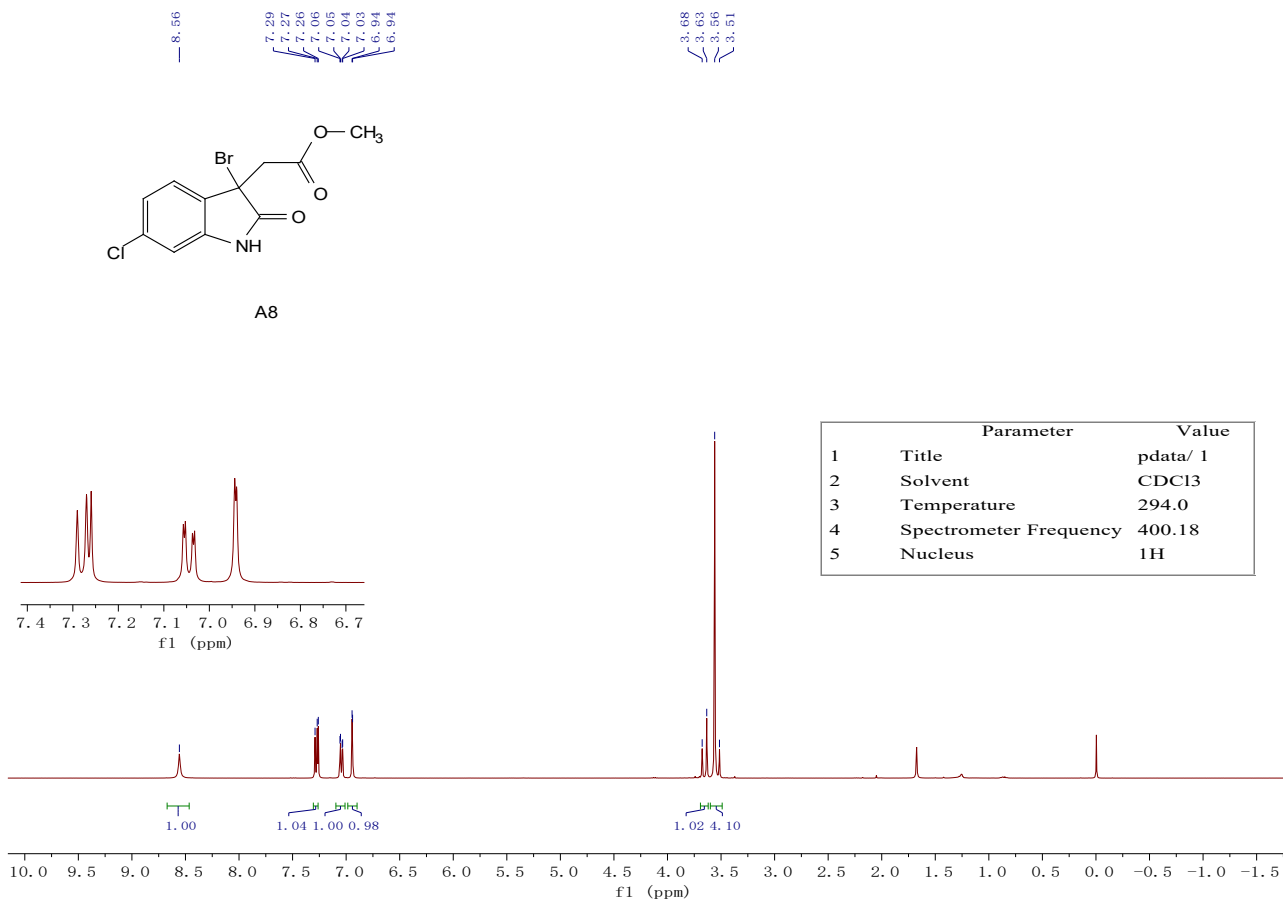
A7



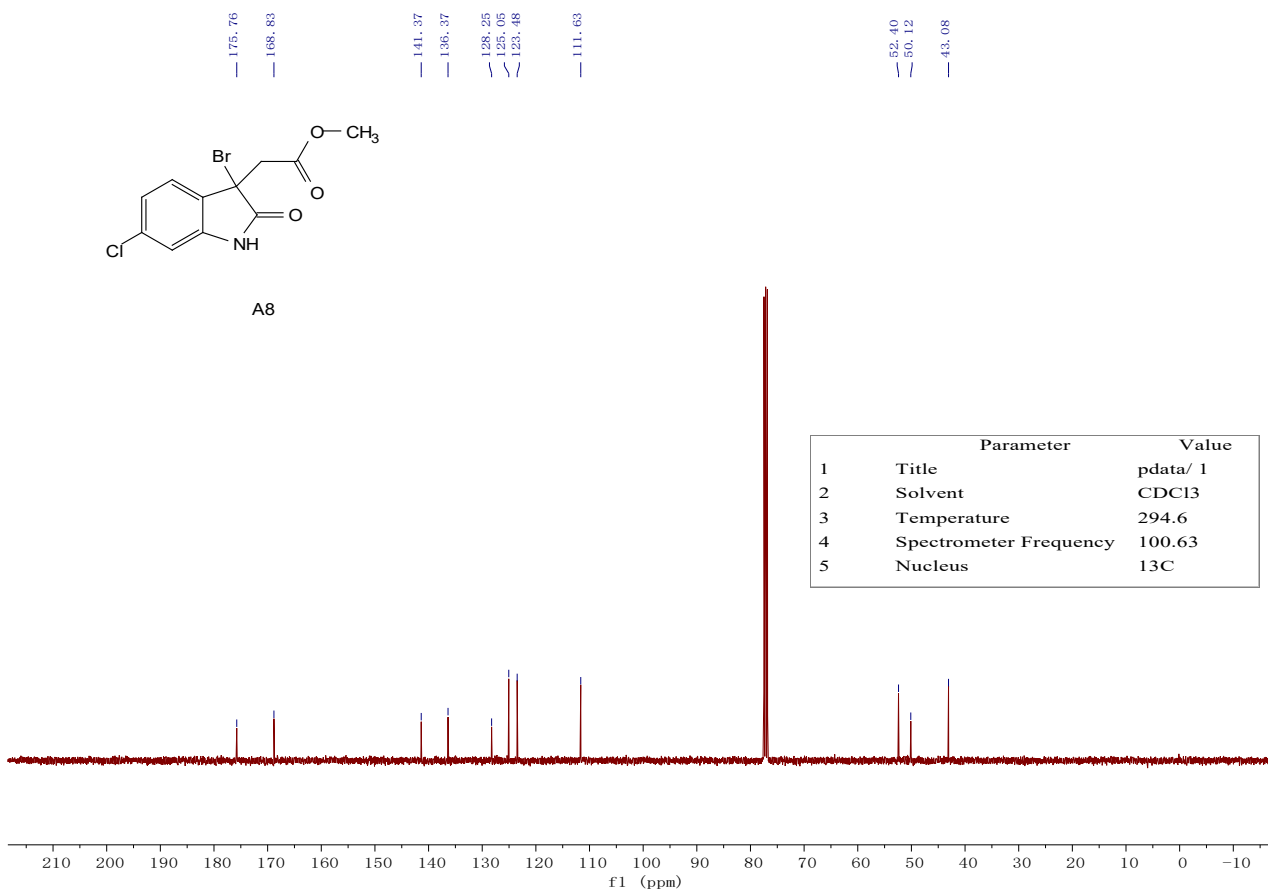


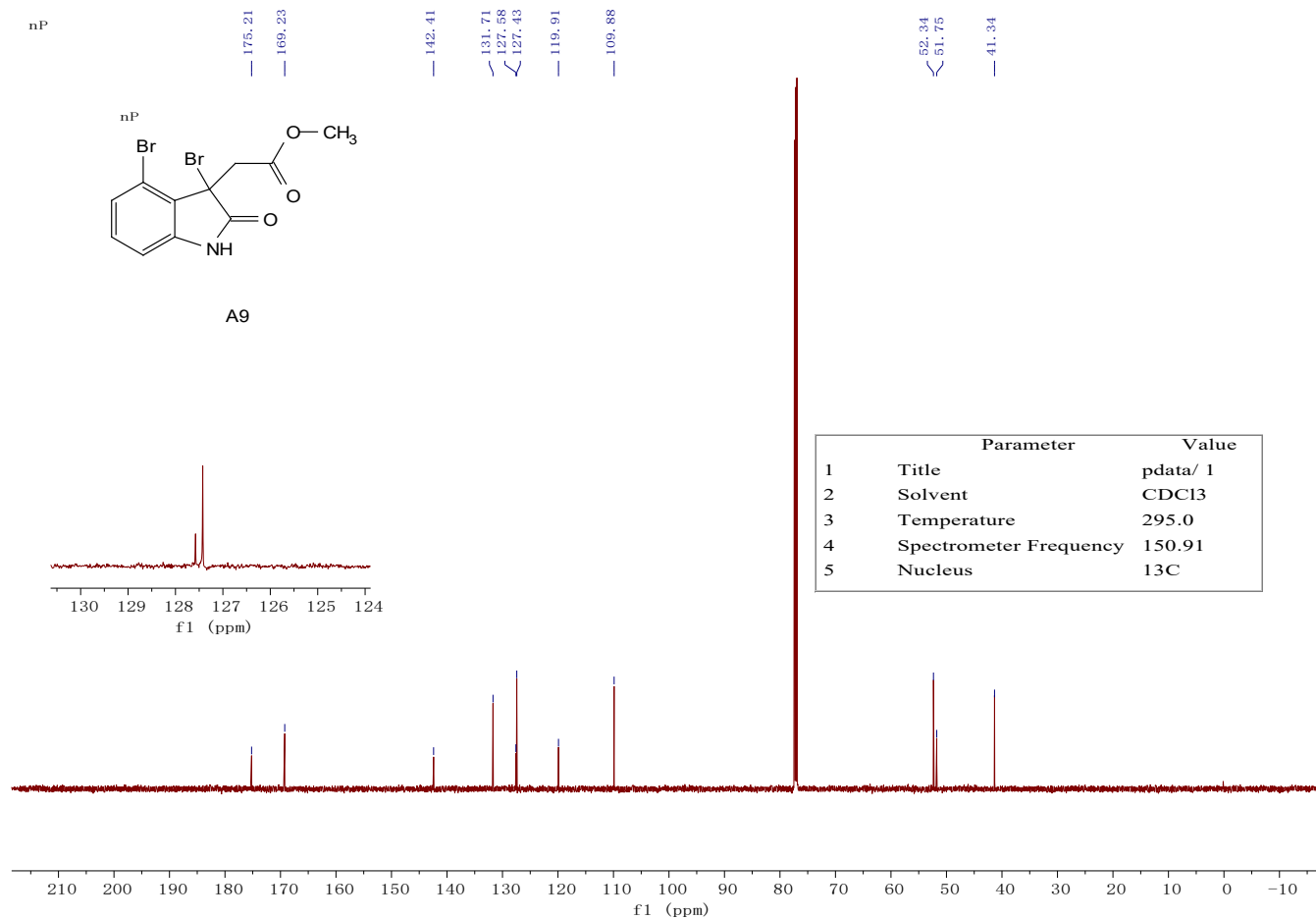
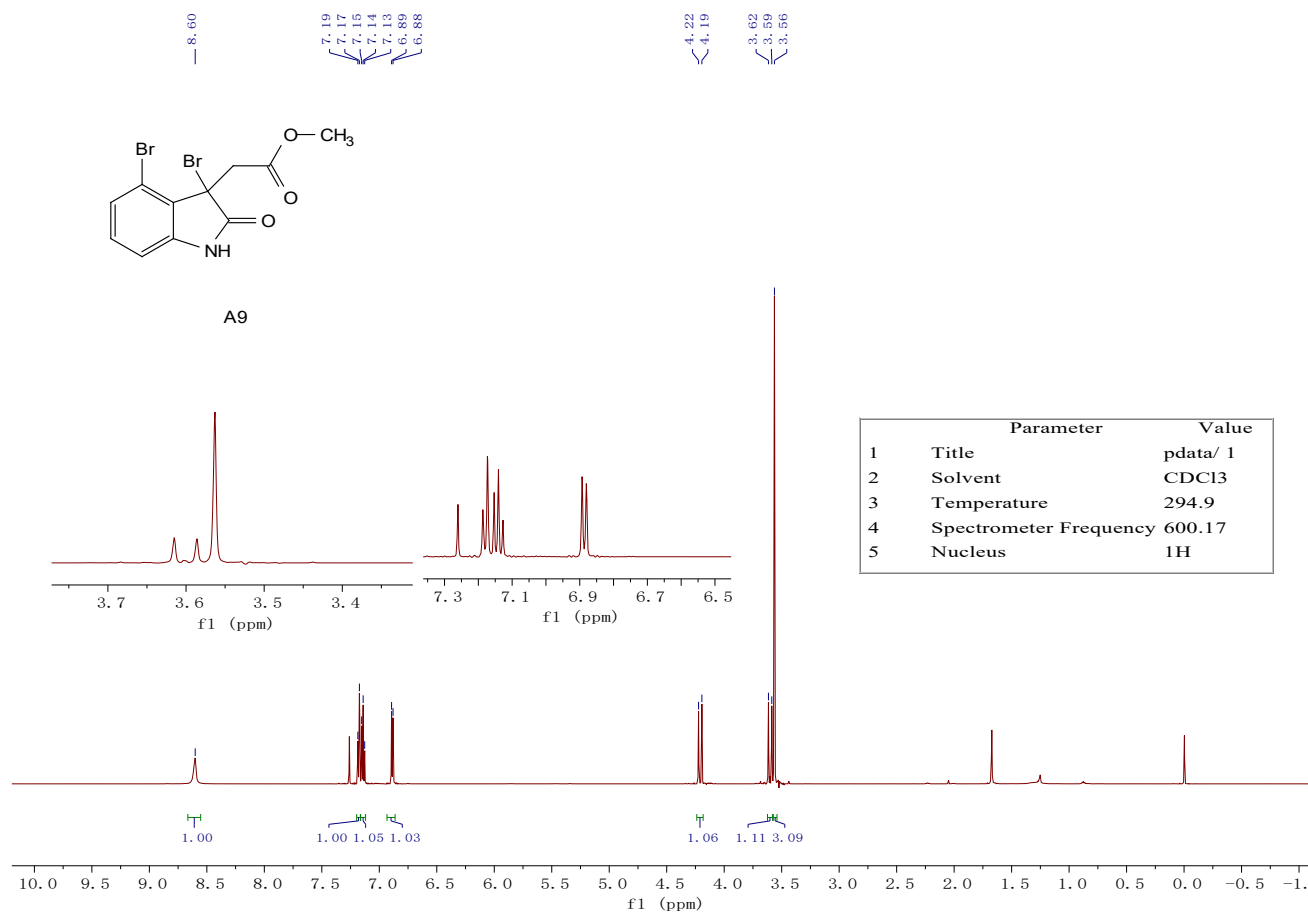


A8

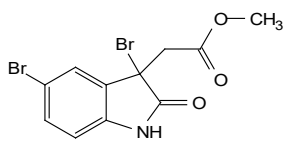


A8

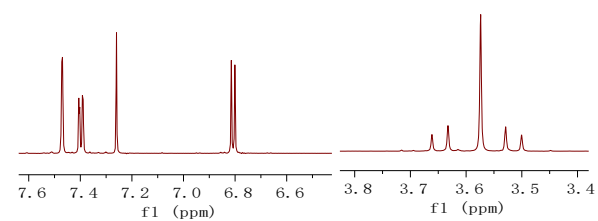




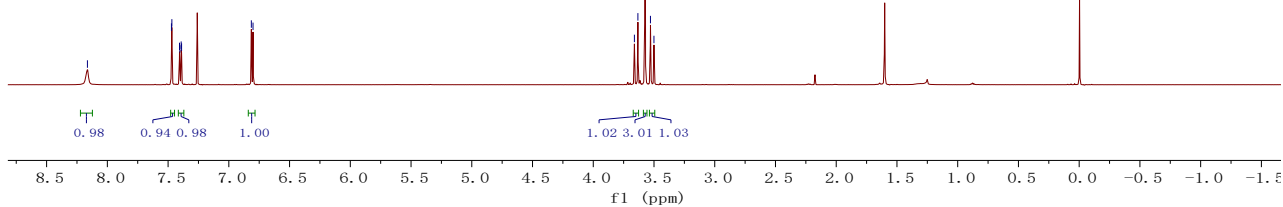
8.16  
7.47  
7.47  
7.41  
7.40  
7.39  
7.39  
6.81  
6.80  
3.66  
3.63  
3.57  
3.53  
3.50



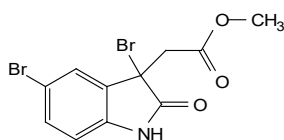
A10



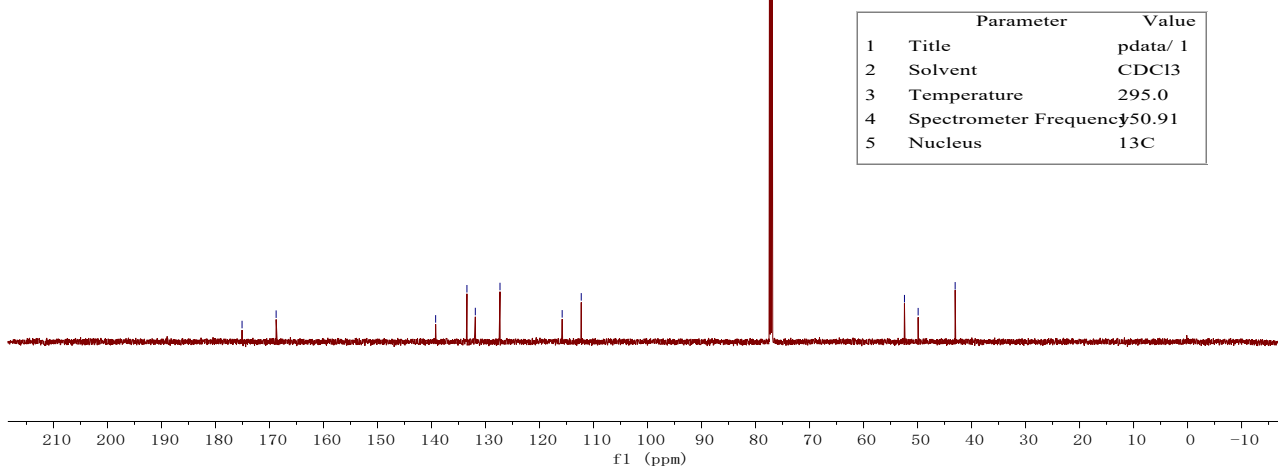
|   | Parameter              | Value    |
|---|------------------------|----------|
| 1 | Title                  | pdata/ 1 |
| 2 | Solvent                | CDCl3    |
| 3 | Temperature            | 295.1    |
| 4 | Spectrometer Frequency | 600.17   |
| 5 | Nucleus                | 1H       |



175.08  
168.77  
139.23  
133.44  
131.89  
127.32  
115.80  
112.27  
52.42  
49.89  
43.03



A10

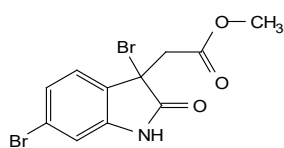


|   | Parameter              | Value    |
|---|------------------------|----------|
| 1 | Title                  | pdata/ 1 |
| 2 | Solvent                | CDCl3    |
| 3 | Temperature            | 295.0    |
| 4 | Spectrometer Frequency | 50.91    |
| 5 | Nucleus                | 13C      |

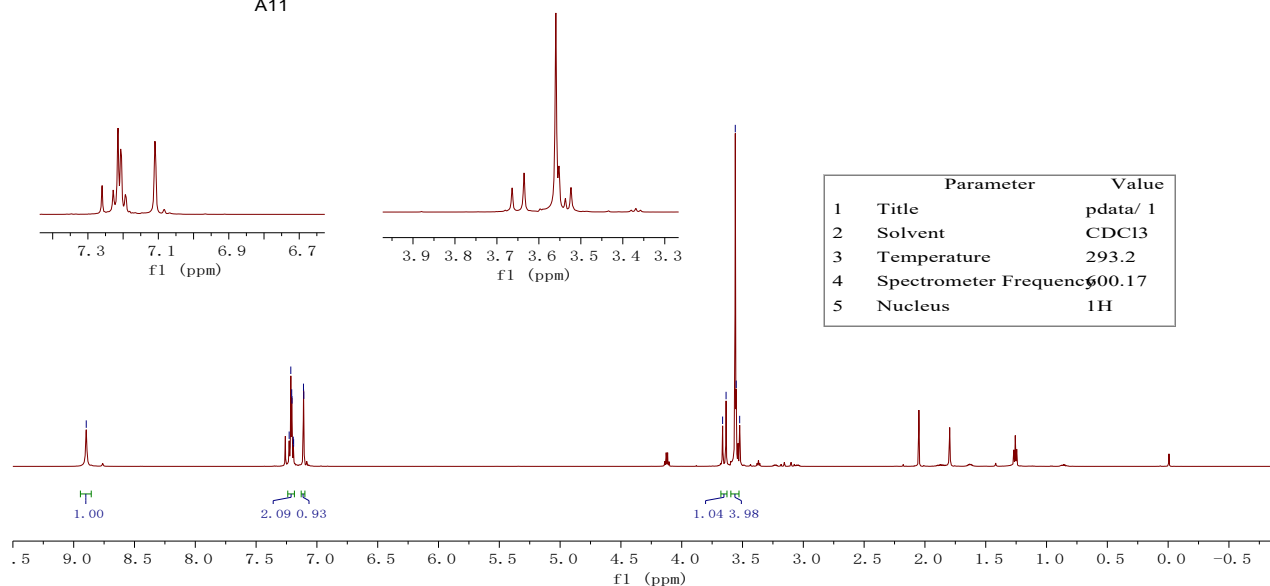
8.90

7.23  
7.21  
7.21  
7.20  
7.19  
7.11  
7.11

3.66  
3.64  
3.56  
3.55  
3.52



A11



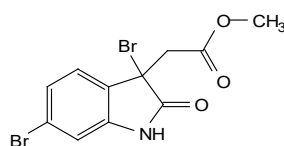
175.91  
168.84

141.56

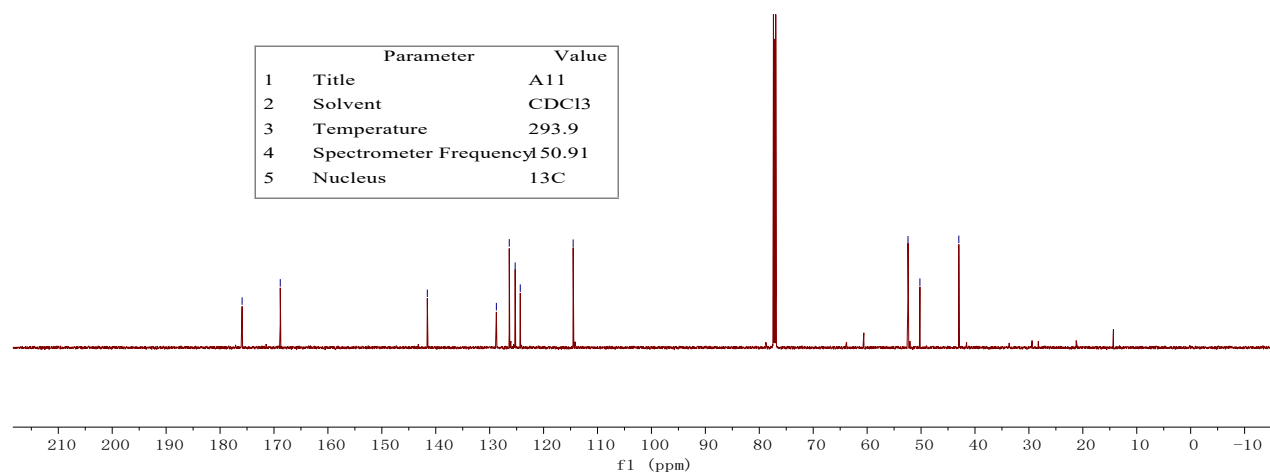
128.76  
126.37  
125.27  
124.33

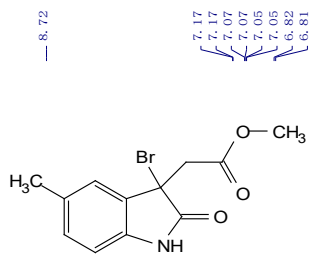
114.52

52.42  
50.21  
43.00



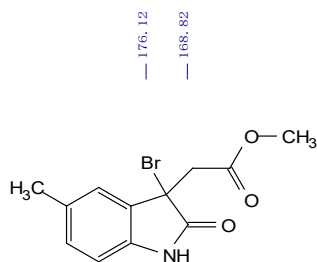
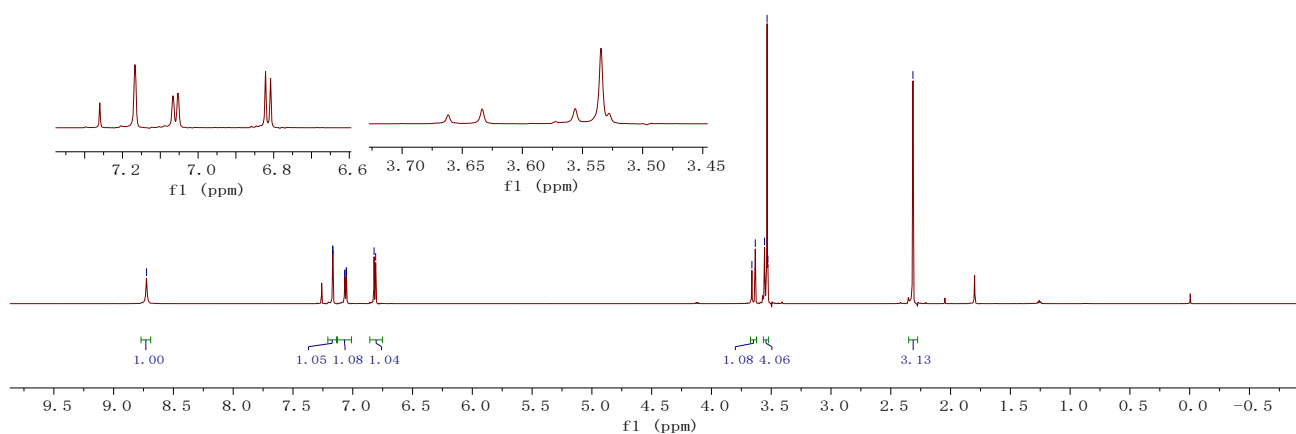
A11





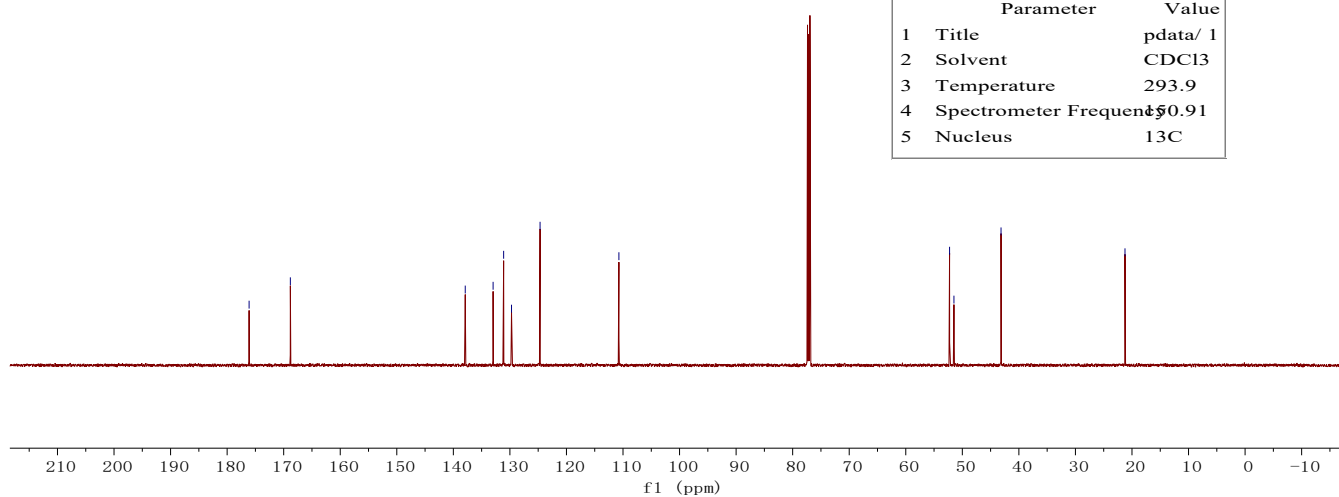
A12

|   | Parameter              | Value          |
|---|------------------------|----------------|
| 1 | Title                  | pdata/ 1       |
| 2 | Solvent                | CDCl3          |
| 3 | Temperature            | 293.0          |
| 4 | Spectrometer Frequency | 600.17         |
| 5 | Nucleus                | <sup>1</sup> H |



A12

|   | Parameter              | Value           |
|---|------------------------|-----------------|
| 1 | Title                  | pdata/ 1        |
| 2 | Solvent                | CDCl3           |
| 3 | Temperature            | 293.9           |
| 4 | Spectrometer Frequency | 150.91          |
| 5 | Nucleus                | <sup>13</sup> C |

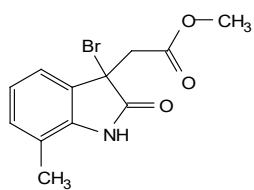


9.37

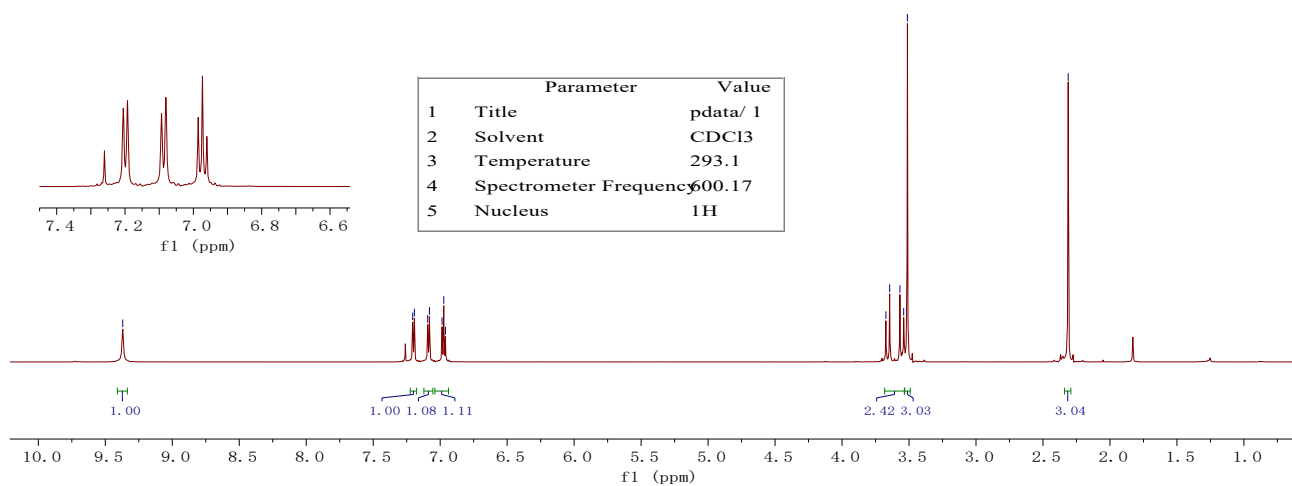
7.21  
7.19  
7.09  
7.08  
6.99  
6.97  
6.96

3.67  
3.64  
3.57  
3.54  
3.51

2.31



A13



176.71

168.78

139.17

131.98

129.34

123.23

121.37

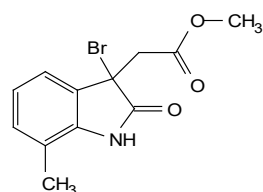
120.47

52.19

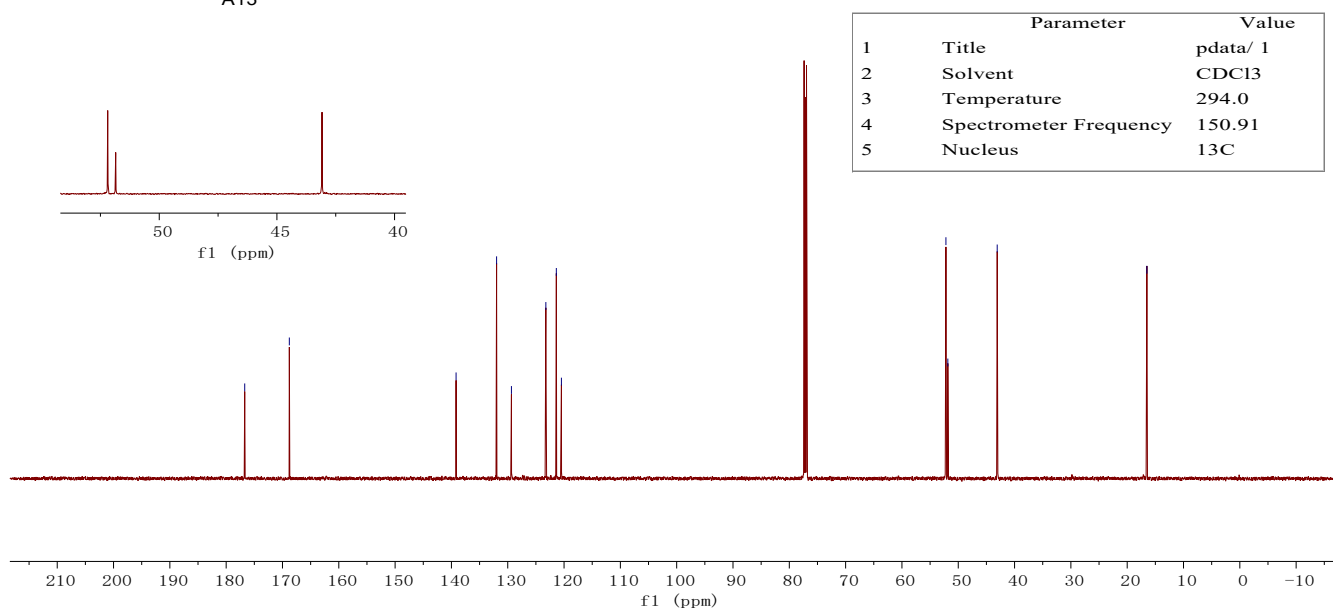
51.85

43.08

16.51



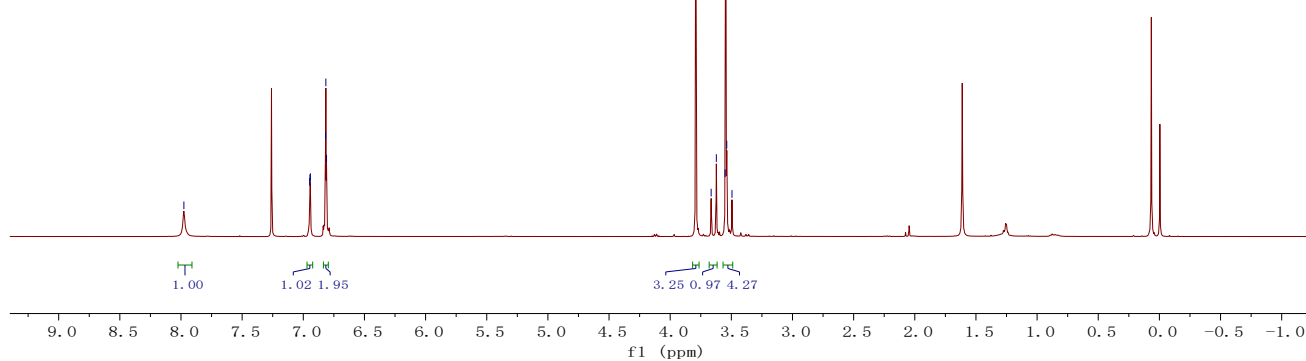
A13




$$\begin{array}{r} 6.95 \\ \swarrow \searrow \\ 6.95 \\ \swarrow \searrow \\ 6.94 \\ \swarrow \searrow \\ 6.82 \\ \swarrow \searrow \\ 6.82 \\ \swarrow \searrow \\ 6.81 \end{array}$$

3.79  
3.67  
3.62  
3.55  
3.55  
3.54  
3.49

|   | Parameter              | Value    |
|---|------------------------|----------|
| 1 | Title                  | pdata/ 1 |
| 2 | Solvent                | CDC13    |
| 3 | Temperature            | 294.1    |
| 4 | Spectrometer Frequency | 400.18   |
| 5 | Nucleus                | 1H       |



168 79

156 32

133. 44

130.92

115.42

111.29

55.94

52.28

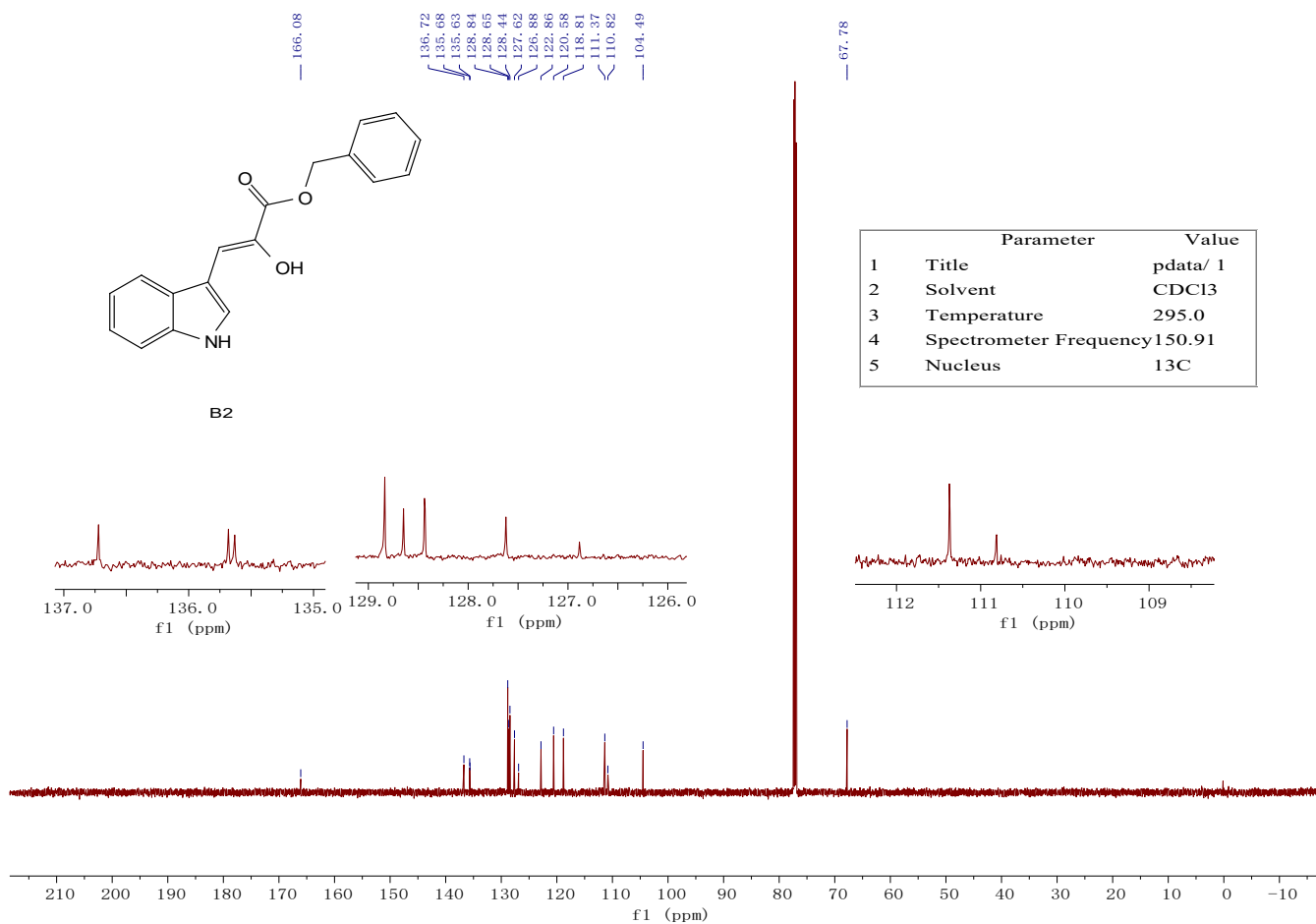
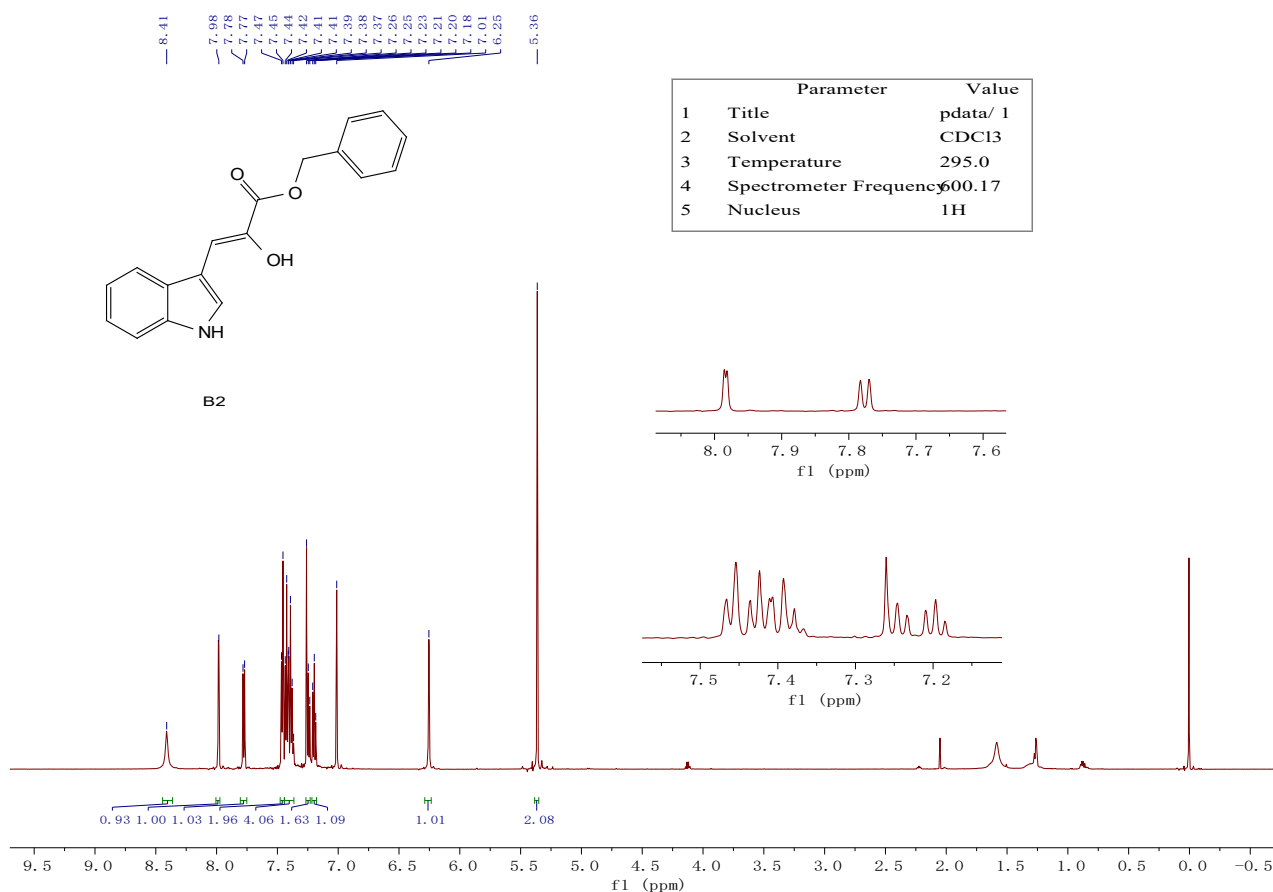
51.28

**Chemical Structure:** COC(=O)CC1(Br)C(=O)Nc2ccc(OC)cc21

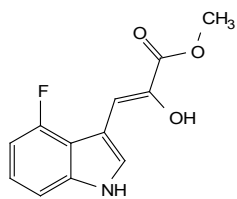
**Figure 1:** <sup>13</sup>C NMR spectrum of compound A14 in CDCl<sub>3</sub>. The spectrum shows peaks at approximately 170 ppm (amide carbonyl), 155 ppm (methoxy carbon), 135 ppm (aromatic carbons), 115 ppm (aromatic carbons), 77 ppm (solvent triplet), 55 ppm (methoxy carbon), and 45 ppm (methyl carbon).

| Parameter                | Value             |
|--------------------------|-------------------|
| 1 Title                  | pdata/ 1          |
| 2 Solvent                | CDCl <sub>3</sub> |
| 3 Temperature            | 294.8             |
| 4 Spectrometer Frequency | 100.63            |
| 5 Nucleus                | 13C               |

|   | Parameter              | Value    |
|---|------------------------|----------|
| 1 | Title                  | pdata/ 1 |
| 2 | Solvent                | CDCl3    |
| 3 | Temperature            | 294.8    |
| 4 | Spectrometer Frequency | 100.63   |
| 5 | Nucleus                | 13C      |

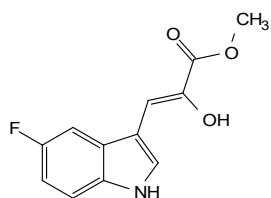
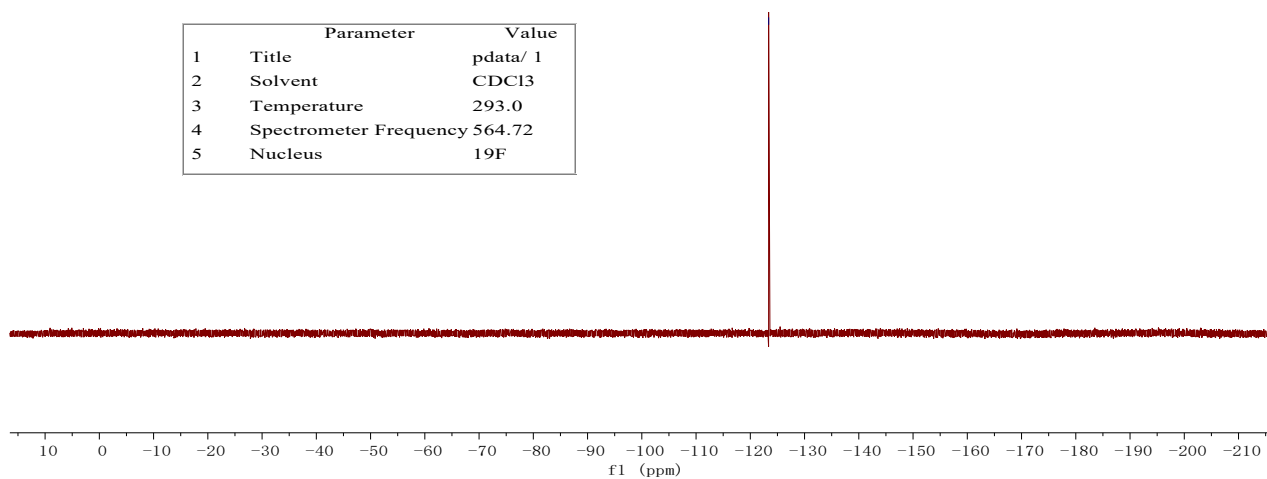




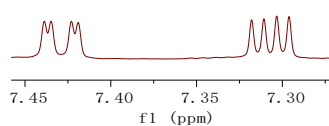
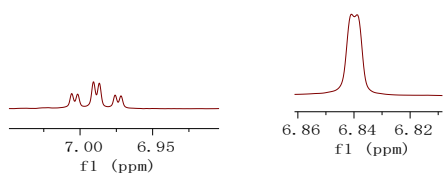


B3

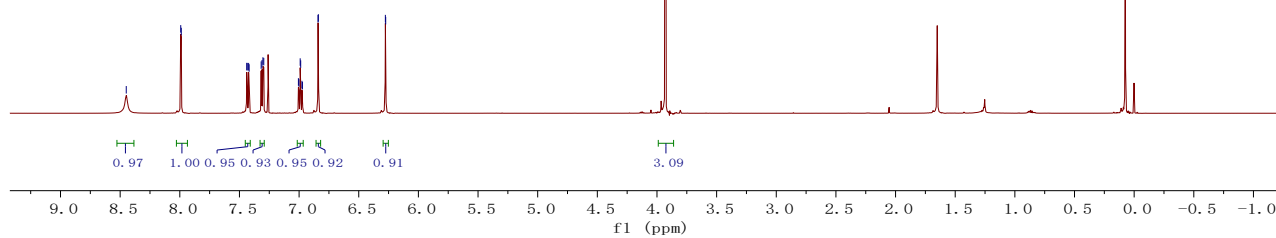
|   | Parameter              | Value    |
|---|------------------------|----------|
| 1 | Title                  | pdata/ 1 |
| 2 | Solvent                | CDCl3    |
| 3 | Temperature            | 293.0    |
| 4 | Spectrometer Frequency | 564.72   |
| 5 | Nucleus                | 19F      |

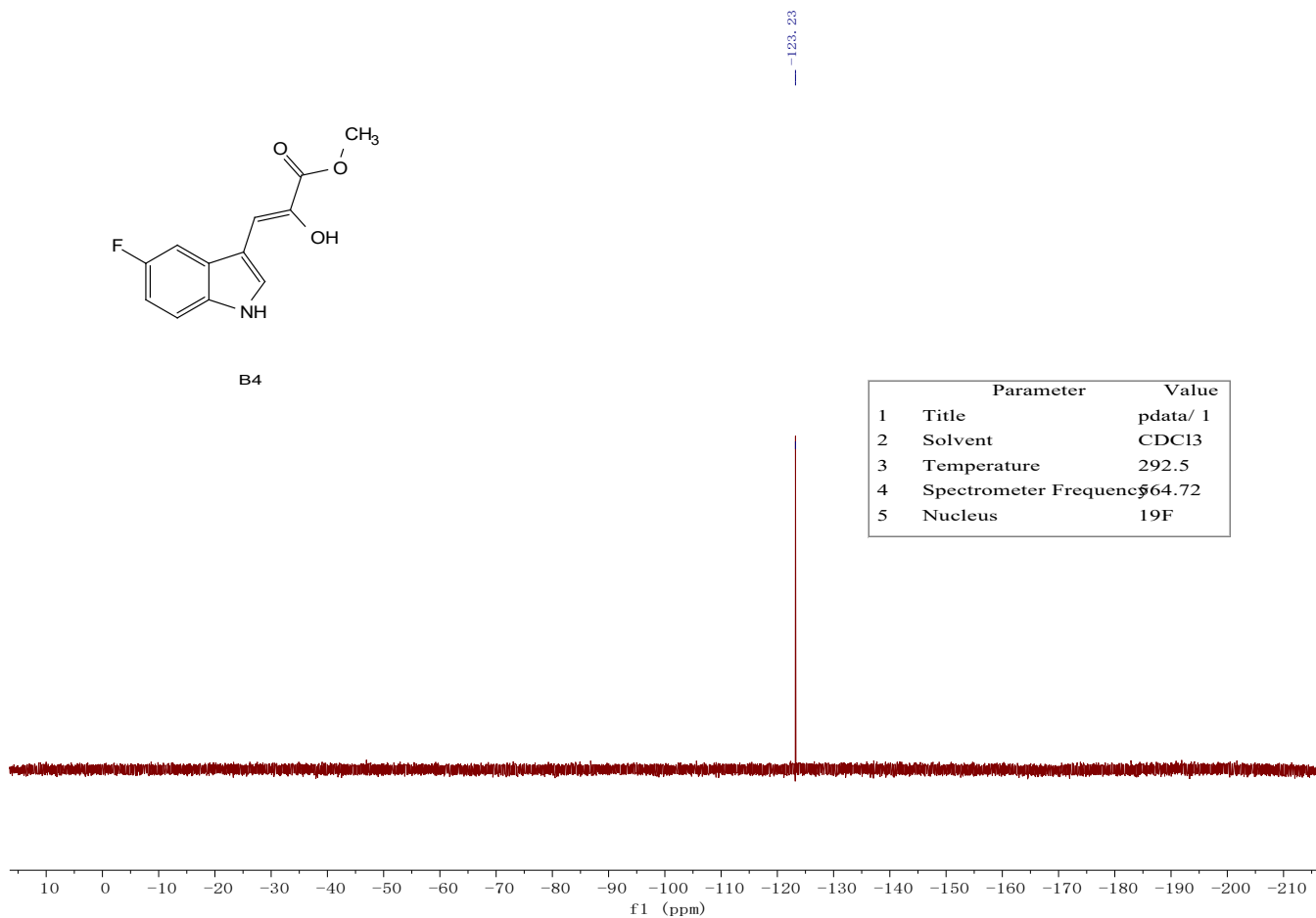
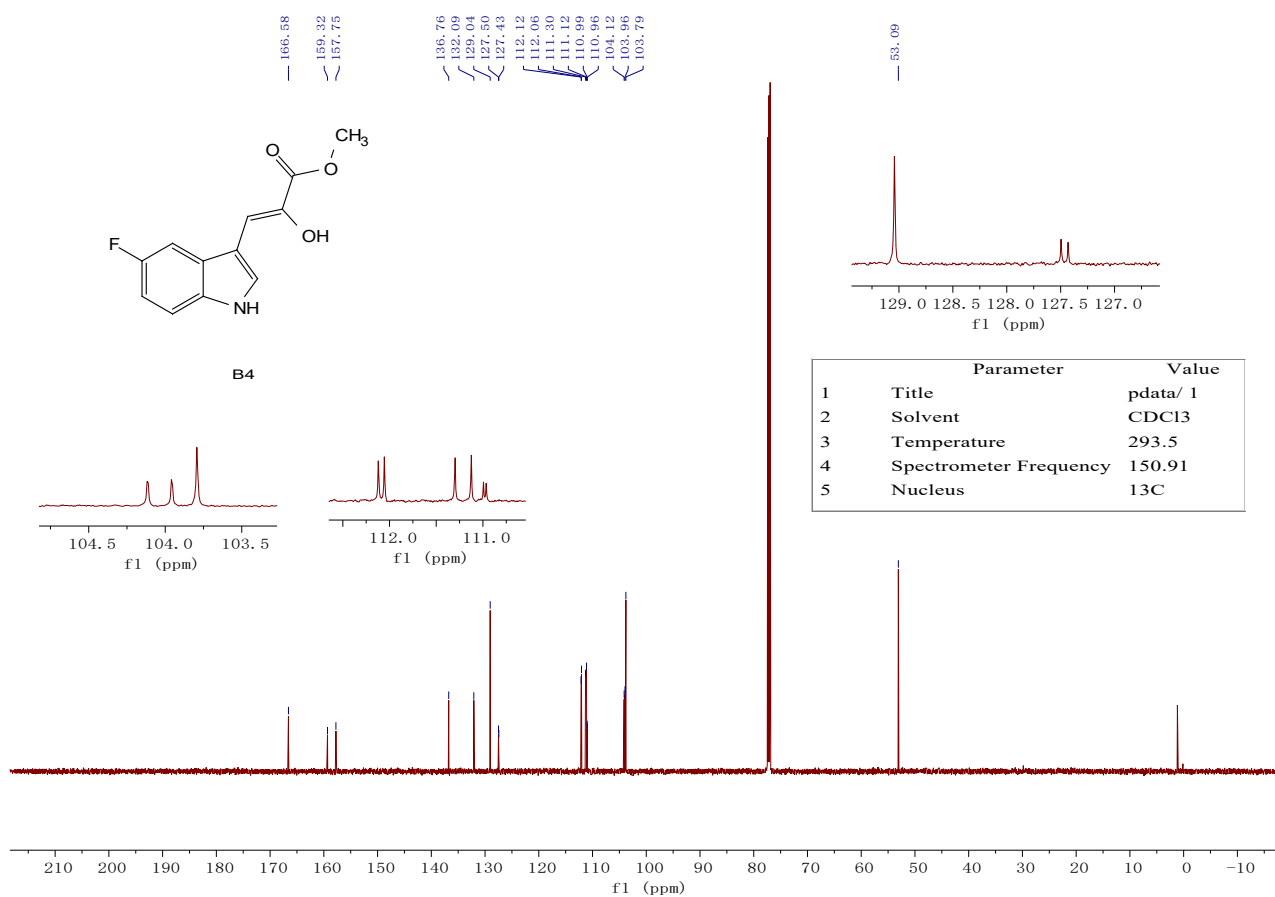


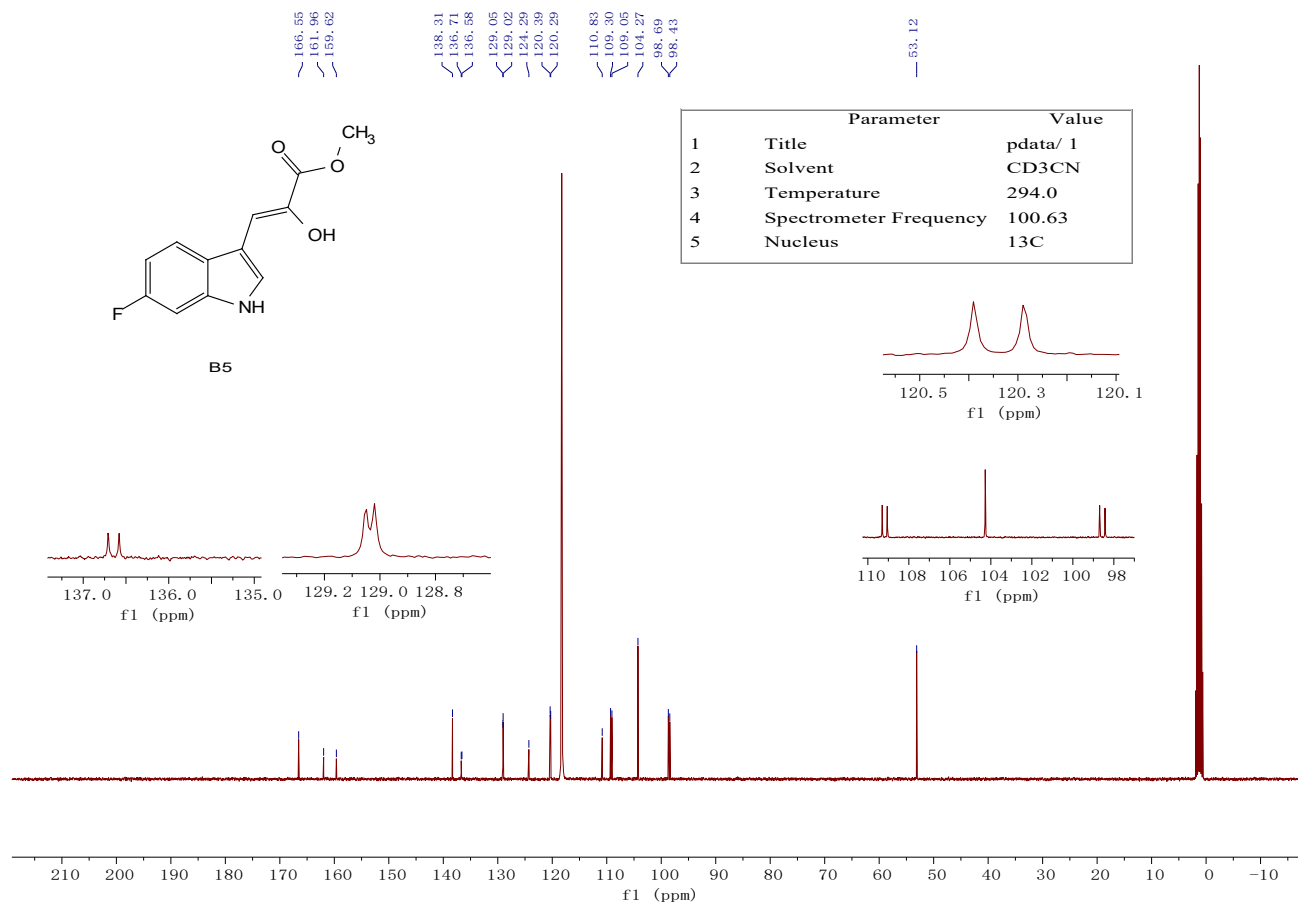
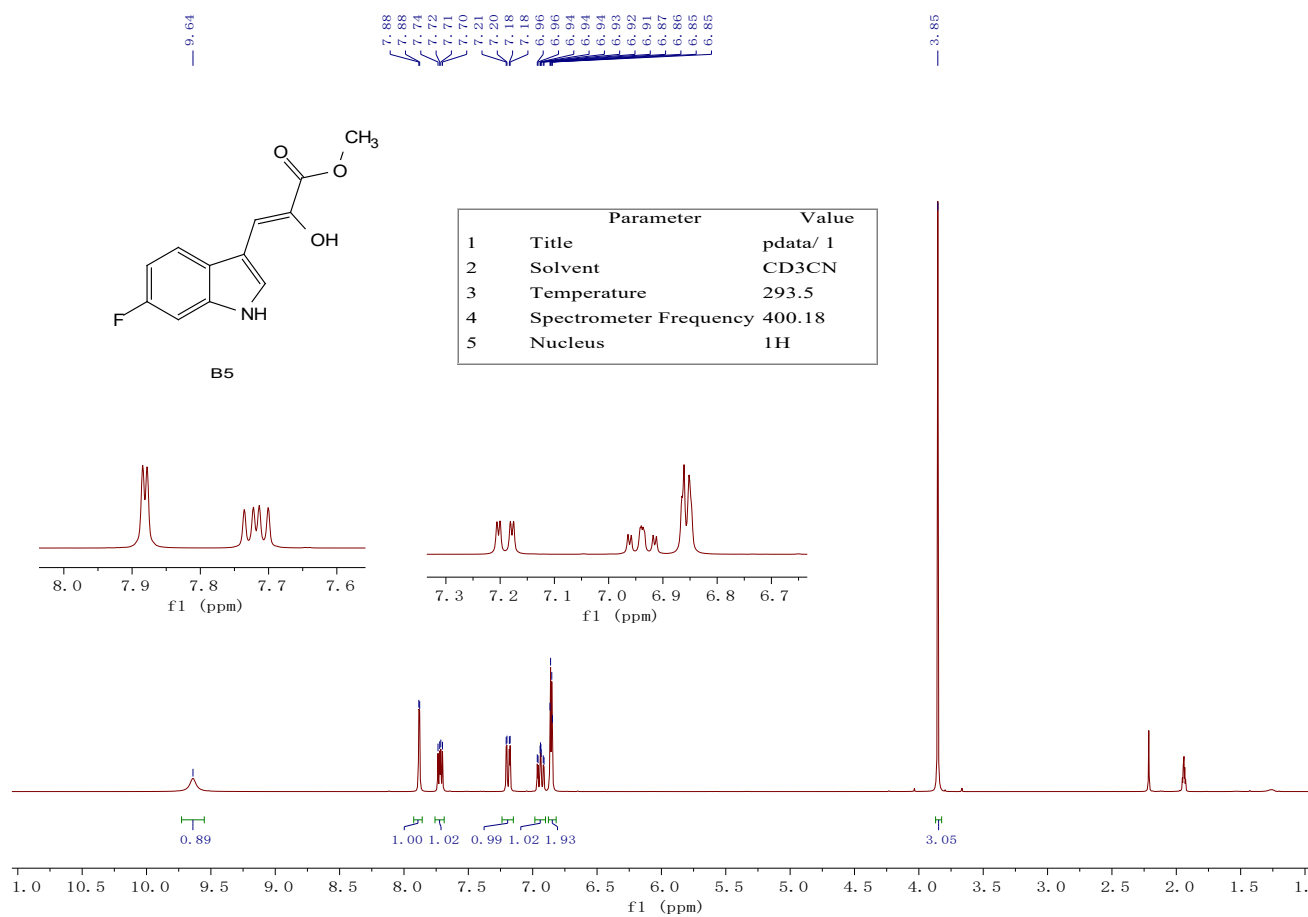
B4

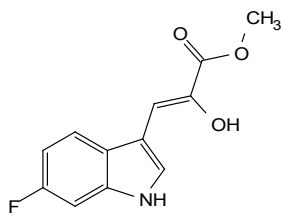


|   | Parameter              | Value    |
|---|------------------------|----------|
| 1 | Title                  | pdata/ 1 |
| 2 | Solvent                | CDCl3    |
| 3 | Temperature            | 292.8    |
| 4 | Spectrometer Frequency | 600.17   |
| 5 | Nucleus                | 1H       |





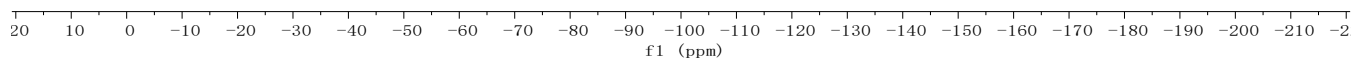




B5

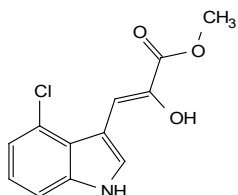
-122.69

|   | Parameter              | Value    |
|---|------------------------|----------|
| 1 | Title                  | pdata/ 1 |
| 2 | Solvent                | CD3CN    |
| 3 | Temperature            | 293.8    |
| 4 | Spectrometer Frequency | 376.55   |
| 5 | Nucleus                | 19F      |



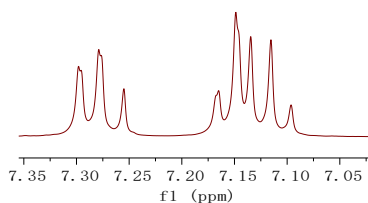
8.56  
8.12  
8.11  
7.76  
7.30  
7.28  
7.28  
7.26  
7.17  
7.16  
7.15  
7.15  
7.13  
7.12  
7.10  
6.35  
6.34

3.92



B6

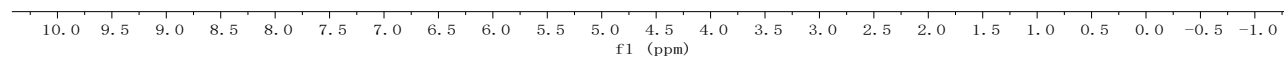
|   | Parameter              | Value  |
|---|------------------------|--------|
| 1 | Title                  | B5     |
| 2 | Solvent                | CDCl3  |
| 3 | Temperature            | 294.6  |
| 4 | Spectrometer Frequency | 400.18 |
| 5 | Nucleus                | 1H     |

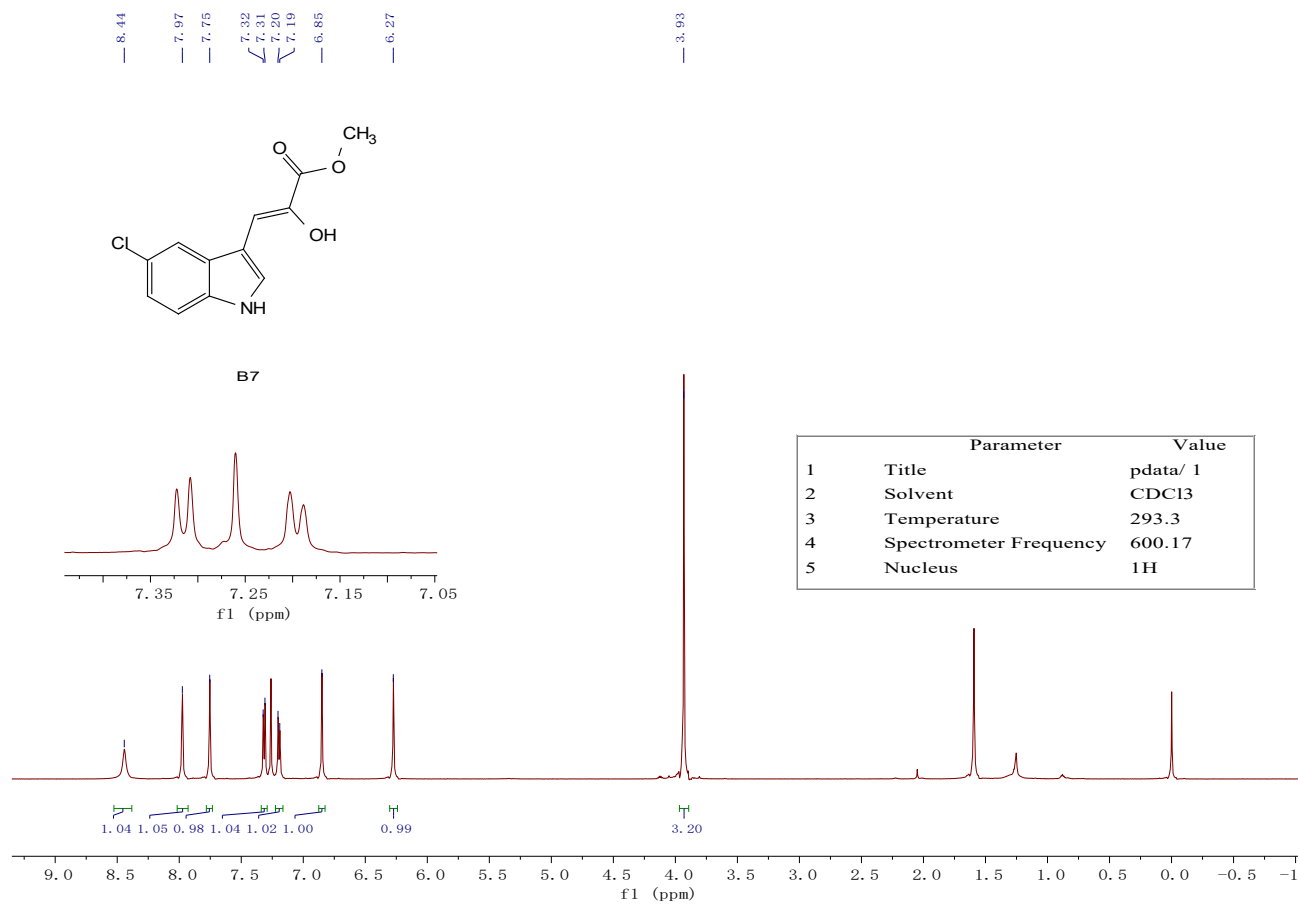
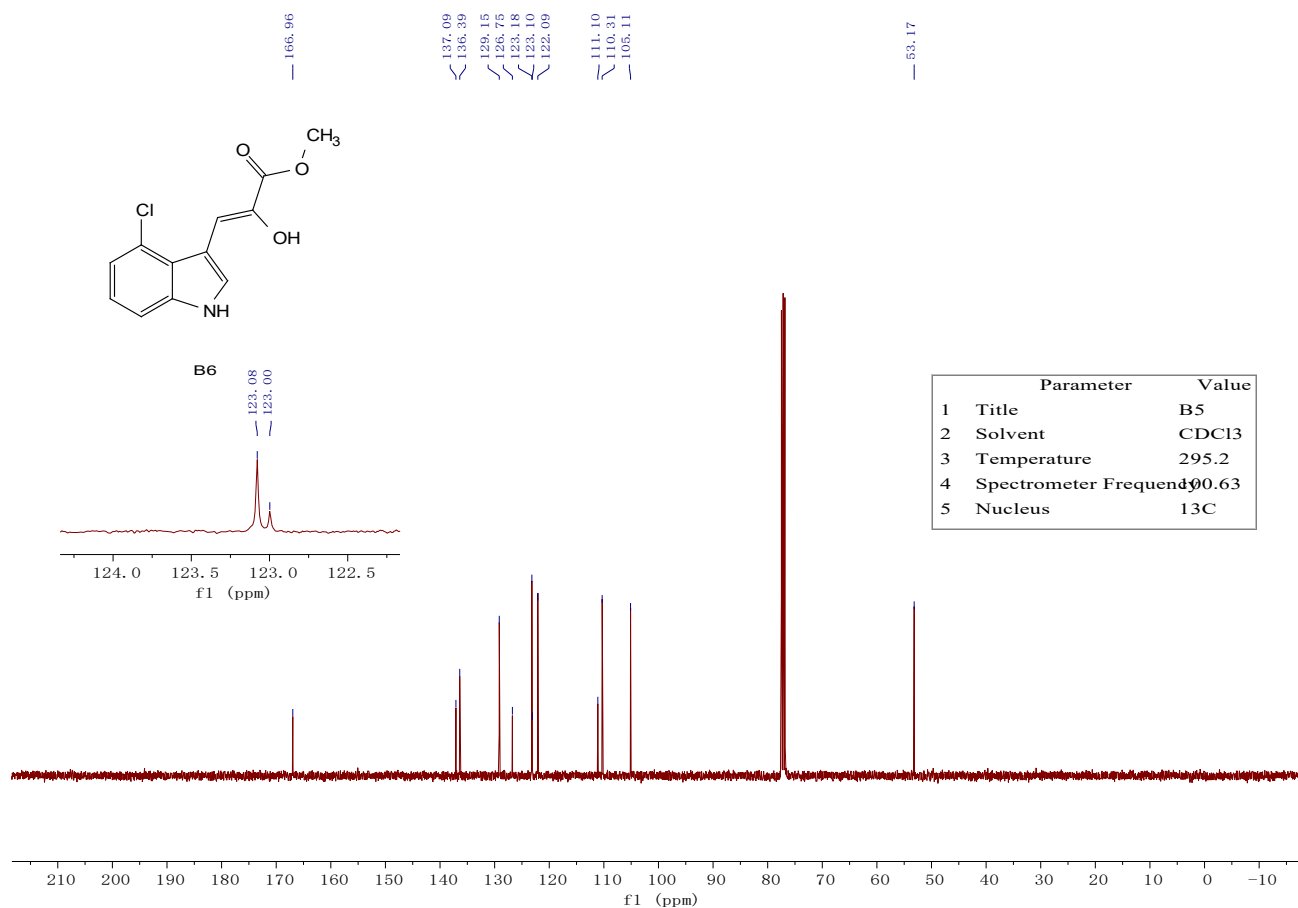


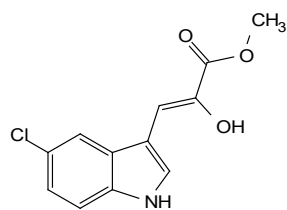
0.98 0.99 0.92 1.18 2.02

1.00

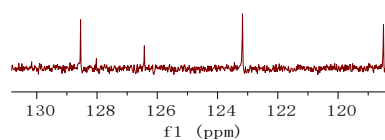
3.15







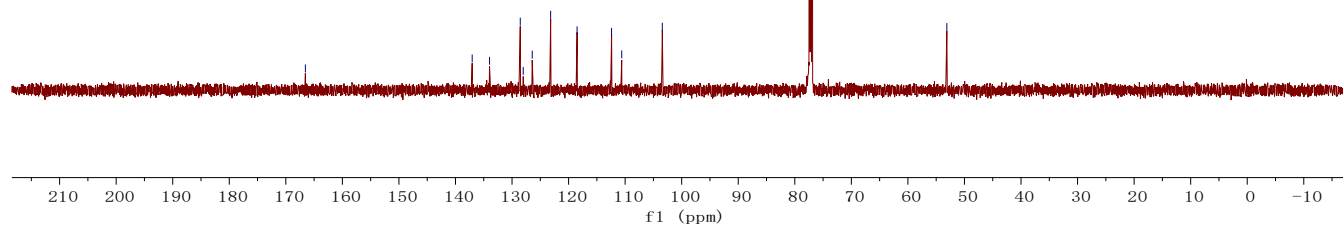
B7



166.54  
137.05  
133.98  
128.55  
128.02  
126.43  
123.17  
118.50  
112.40  
110.61  
103.43

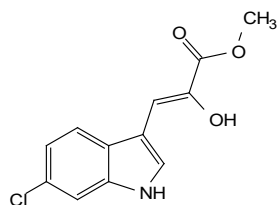
53.12

|   | Parameter              | Value  |
|---|------------------------|--------|
| 1 | Title                  | B6     |
| 2 | Solvent                | CDCl3  |
| 3 | Temperature            | 294.1  |
| 4 | Spectrometer Frequency | 150.91 |
| 5 | Nucleus                | 13C    |

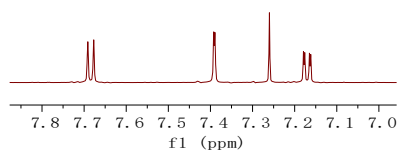


8.41  
7.95  
7.94  
7.69  
7.68  
7.39  
7.39  
7.18  
7.18  
7.16  
7.16  
6.88  
6.87  
6.28  
6.28

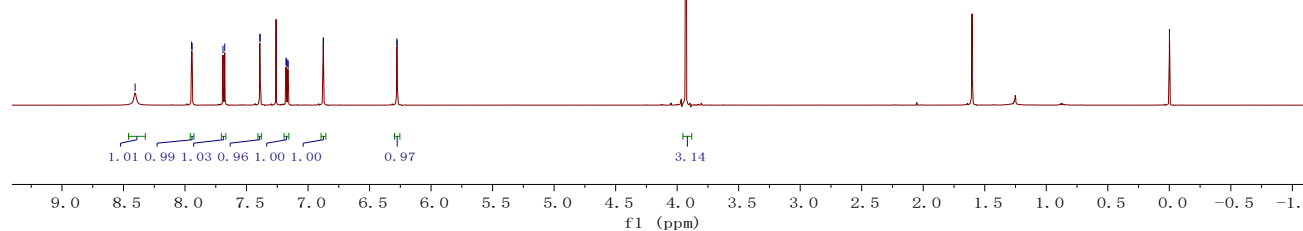
3.93

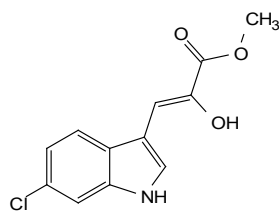


B8

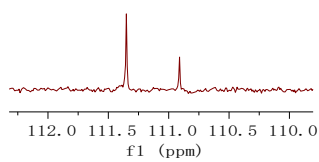


|   | Parameter              | Value    |
|---|------------------------|----------|
| 1 | Title                  | pdata/ 1 |
| 2 | Solvent                | CDCl3    |
| 3 | Temperature            | 292.5    |
| 4 | Spectrometer Frequency | 600.17   |
| 5 | Nucleus                | 1H       |





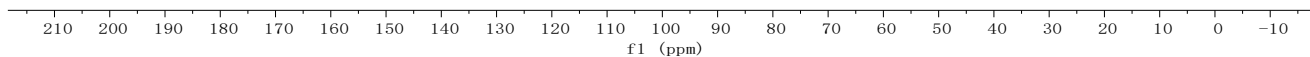
B8



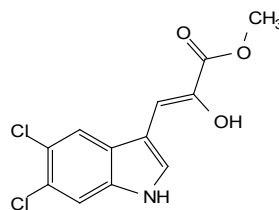
166.51  
137.13  
135.96  
128.72  
127.90  
125.48  
121.29  
119.71  
111.35  
110.91  
103.52

53.11

|   | Parameter              | Value    |
|---|------------------------|----------|
| 1 | Title                  | pdata/ 1 |
| 2 | Solvent                | CDCl3    |
| 3 | Temperature            | 293.0    |
| 4 | Spectrometer Frequency | 150.91   |
| 5 | Nucleus                | 13C      |

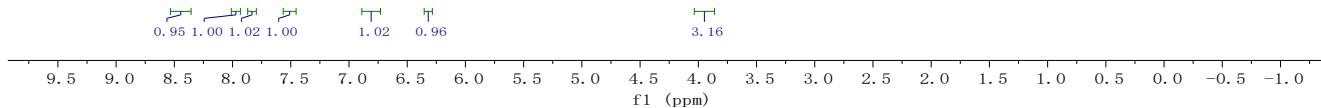


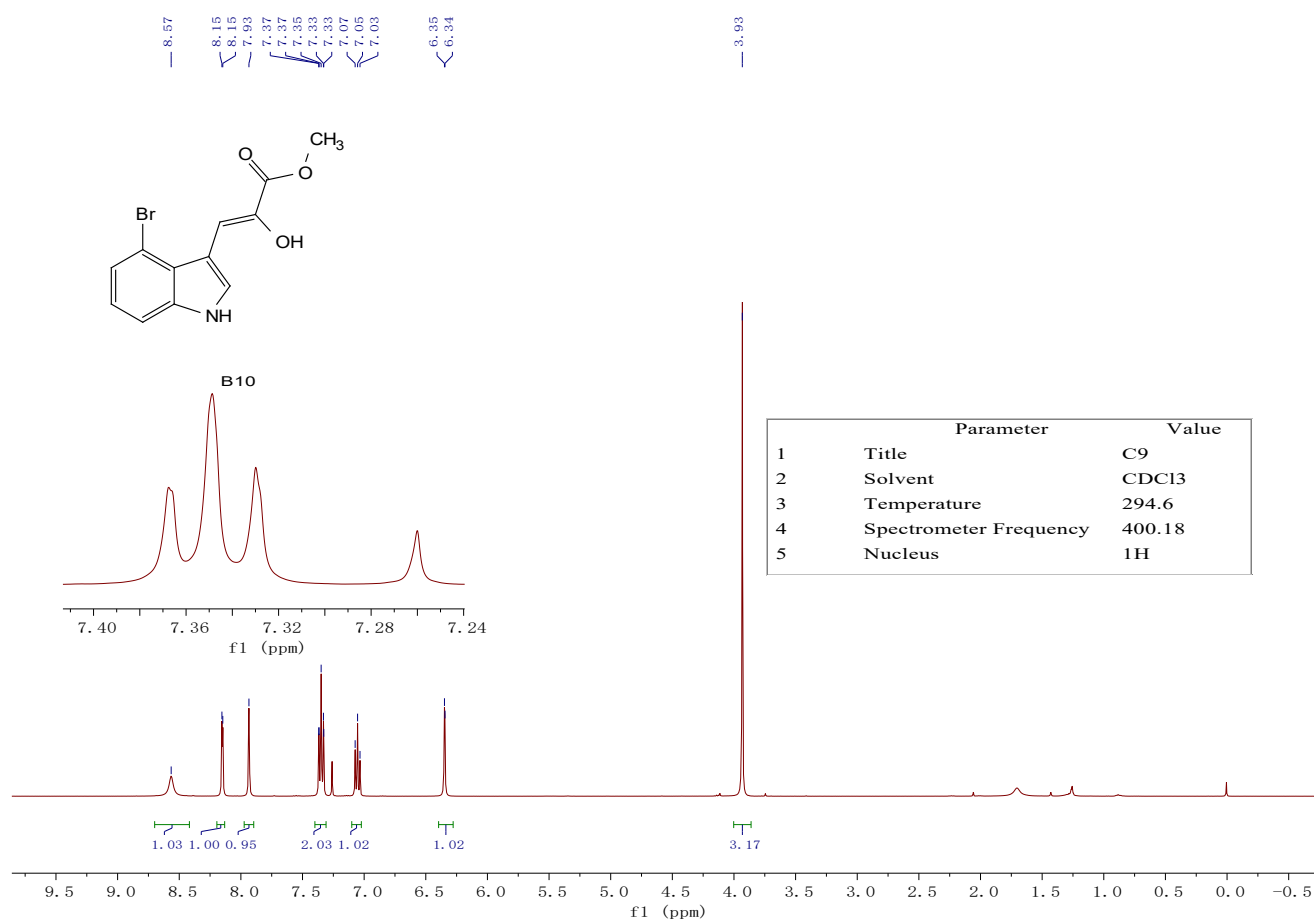
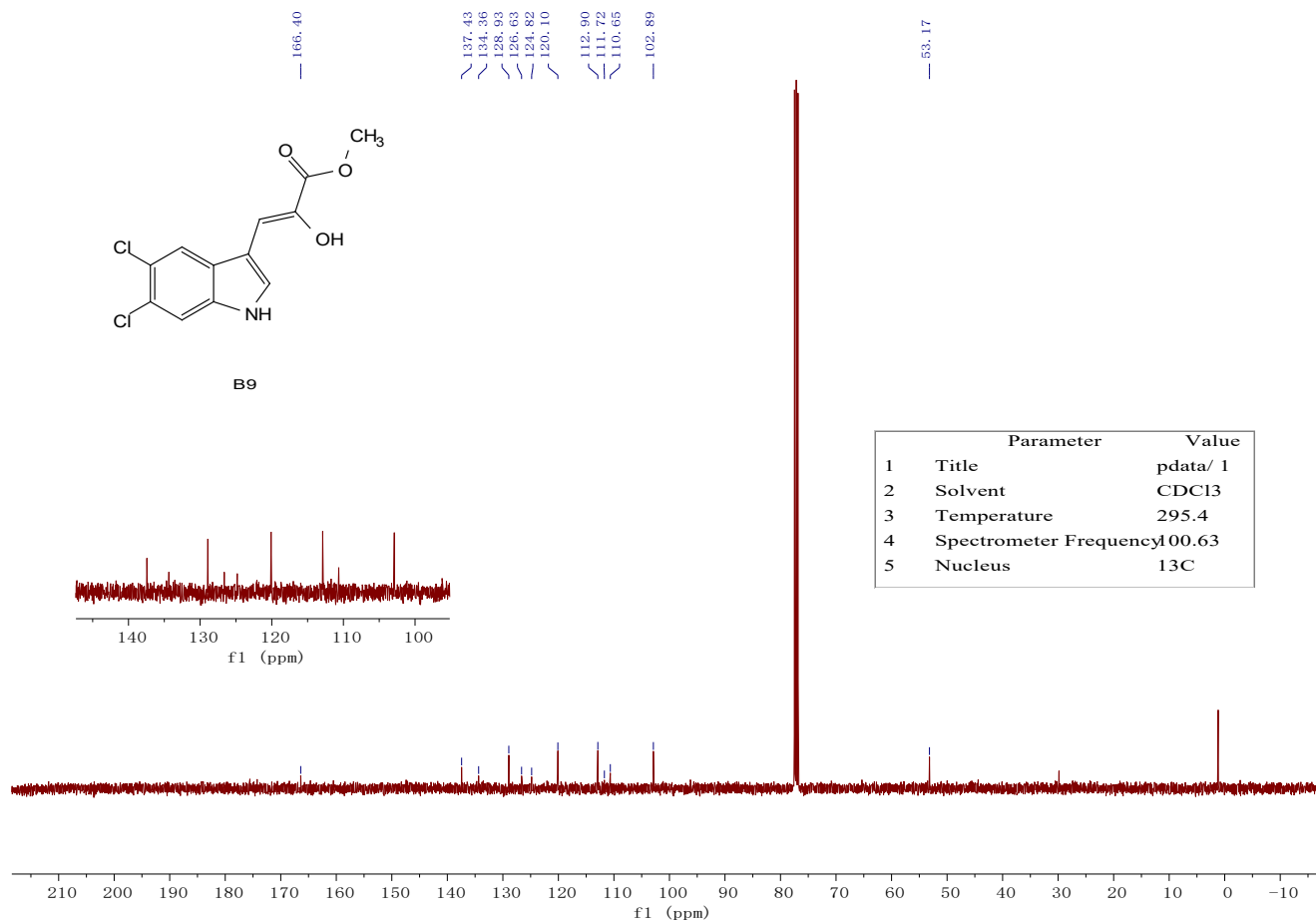
8.43  
7.96  
7.96  
7.85  
7.51  
6.79  
6.79  
6.31  
6.30  
3.93

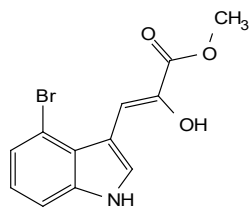


B9

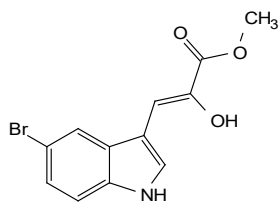
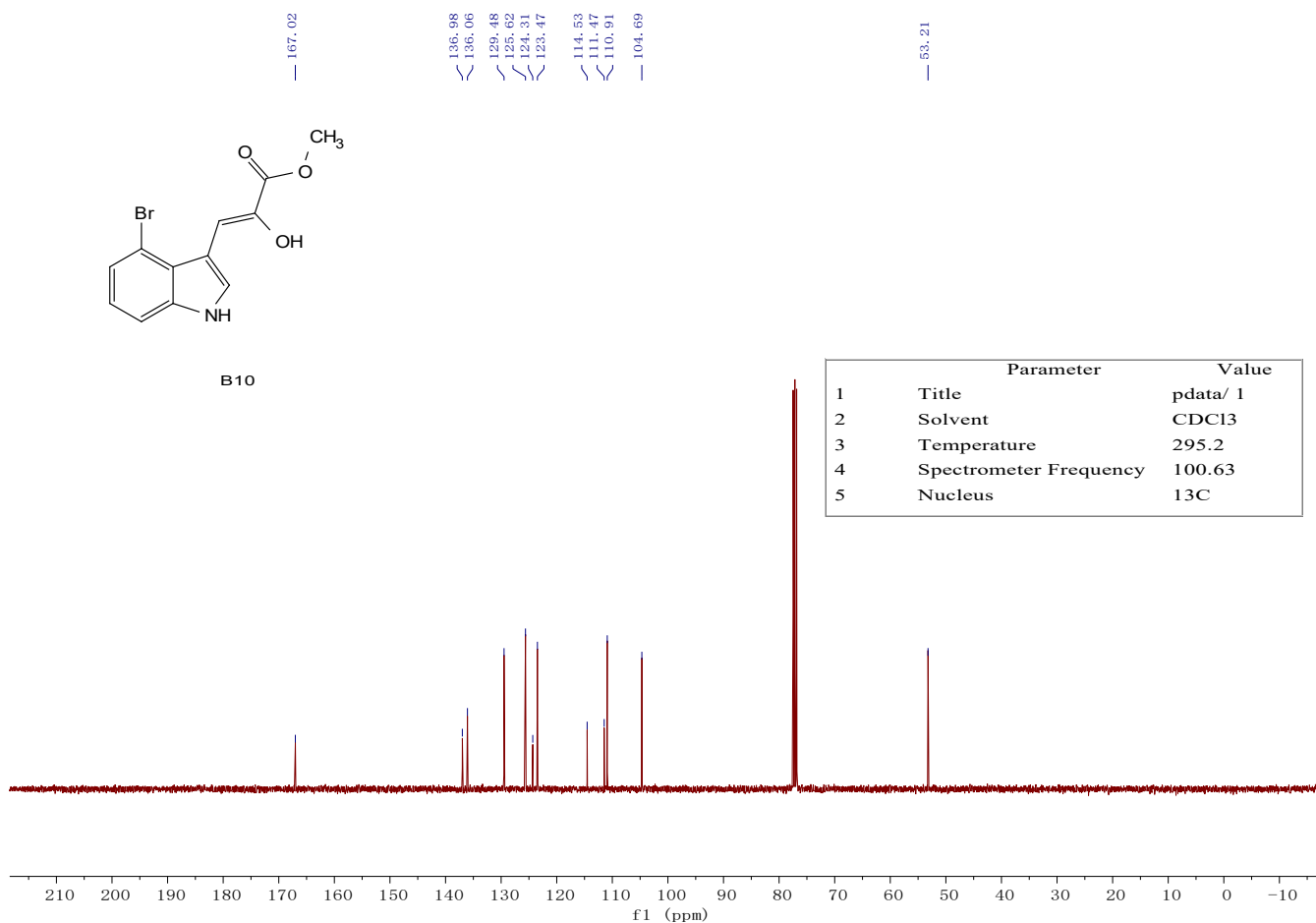
|   | Parameter              | Value    |
|---|------------------------|----------|
| 1 | Title                  | pdata/ 1 |
| 2 | Solvent                | CDCl3    |
| 3 | Temperature            | 294.8    |
| 4 | Spectrometer Frequency | 400.18   |
| 5 | Nucleus                | 1H       |



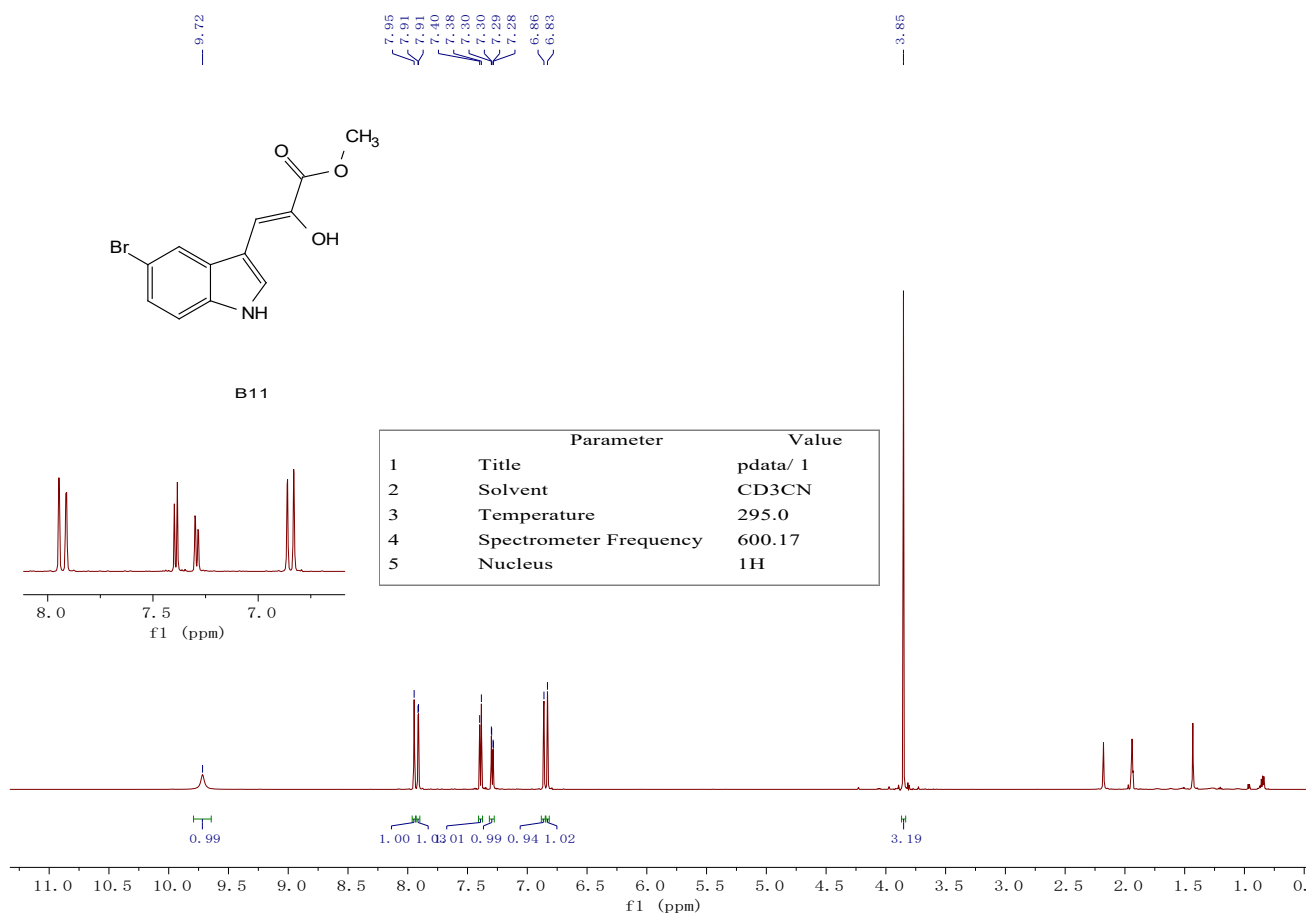


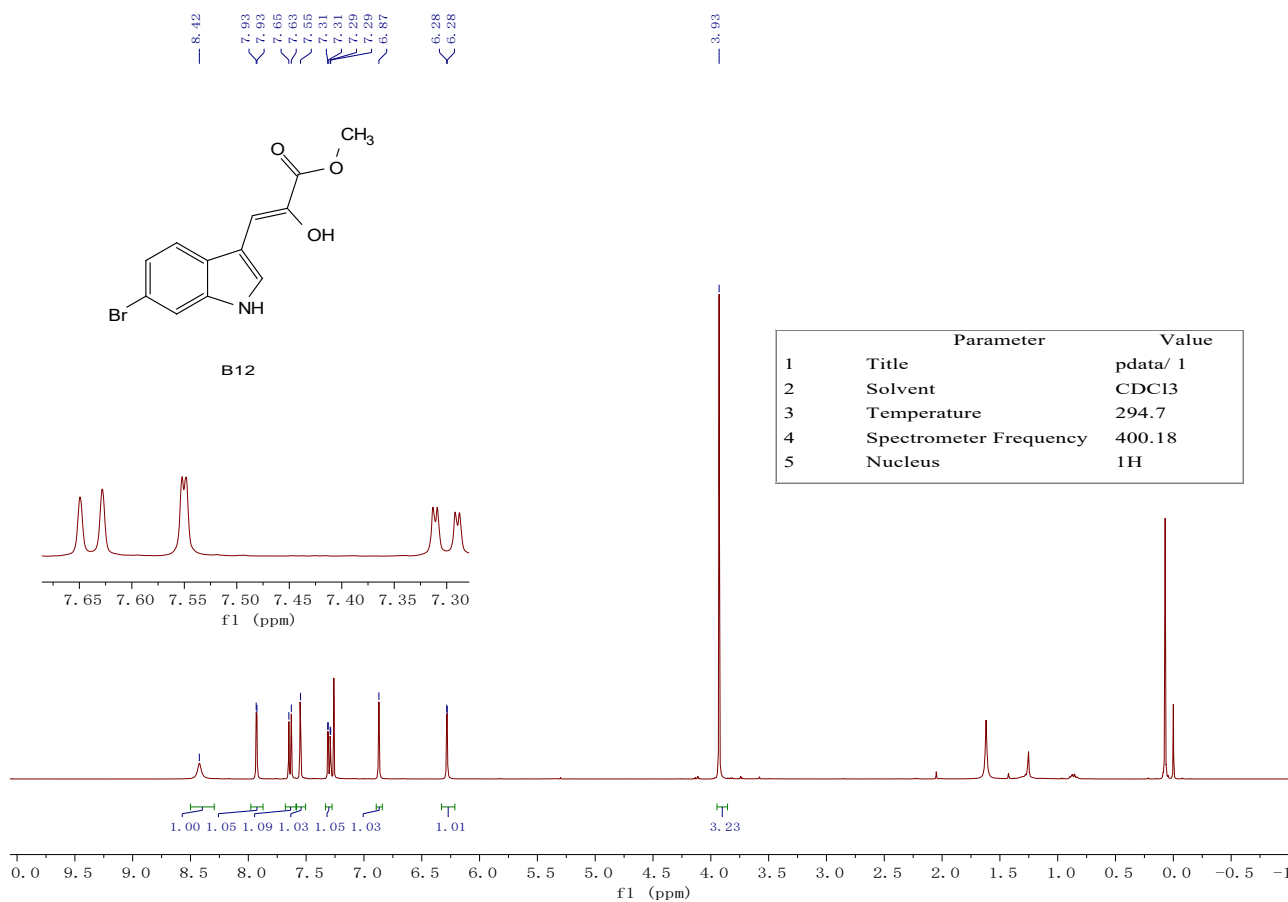
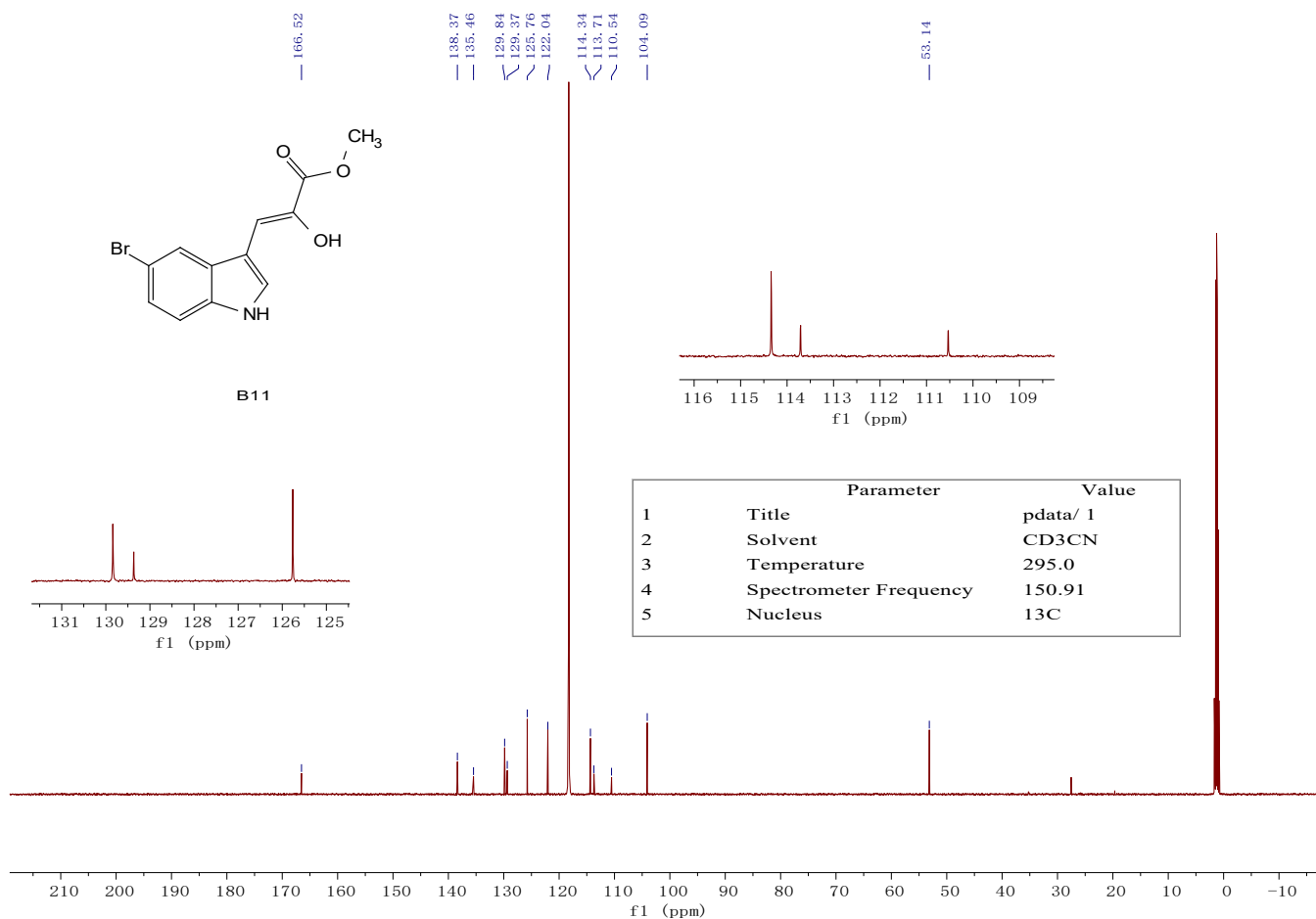


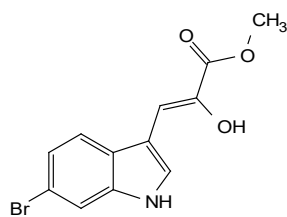
B10



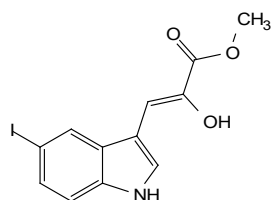
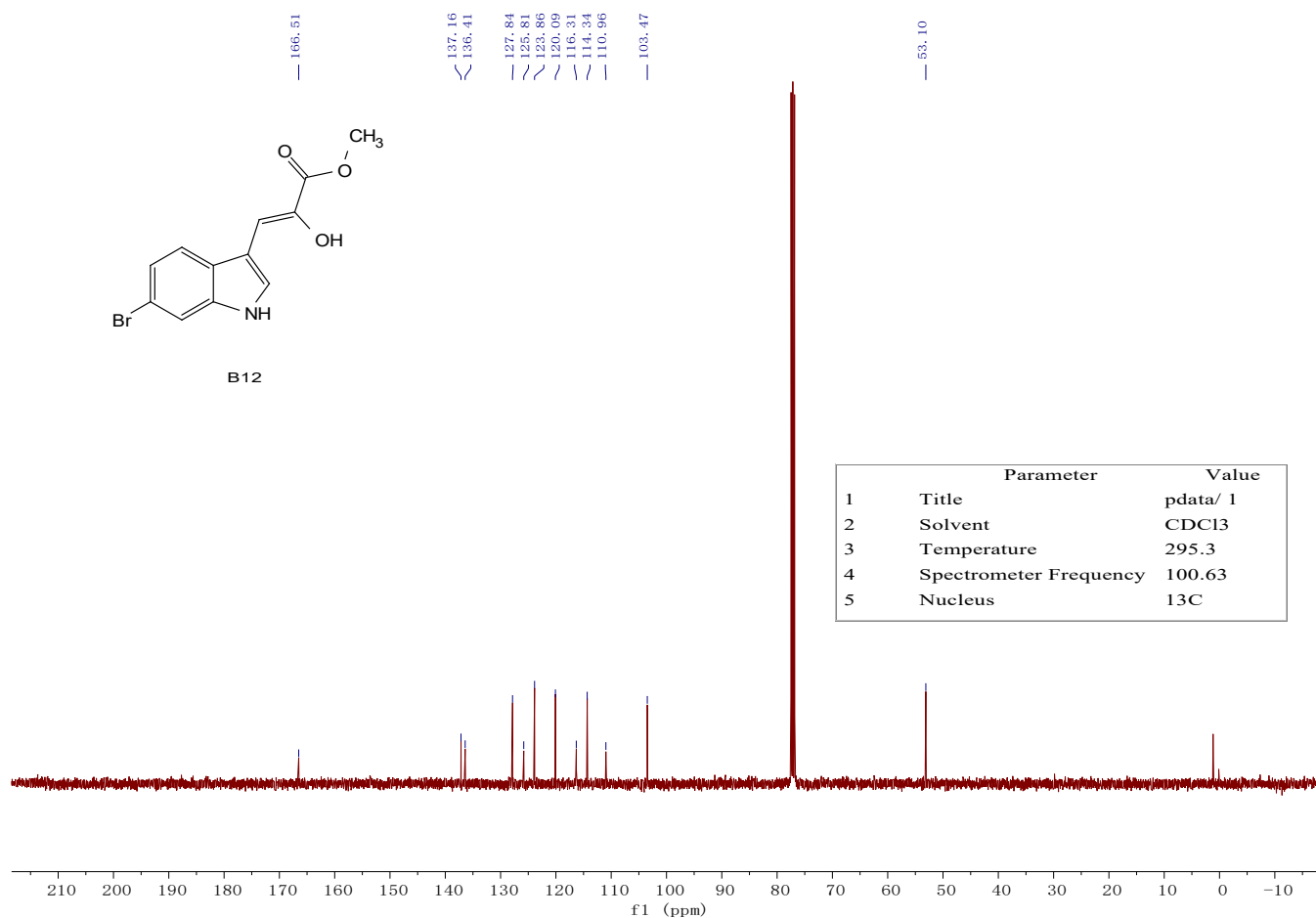
B11



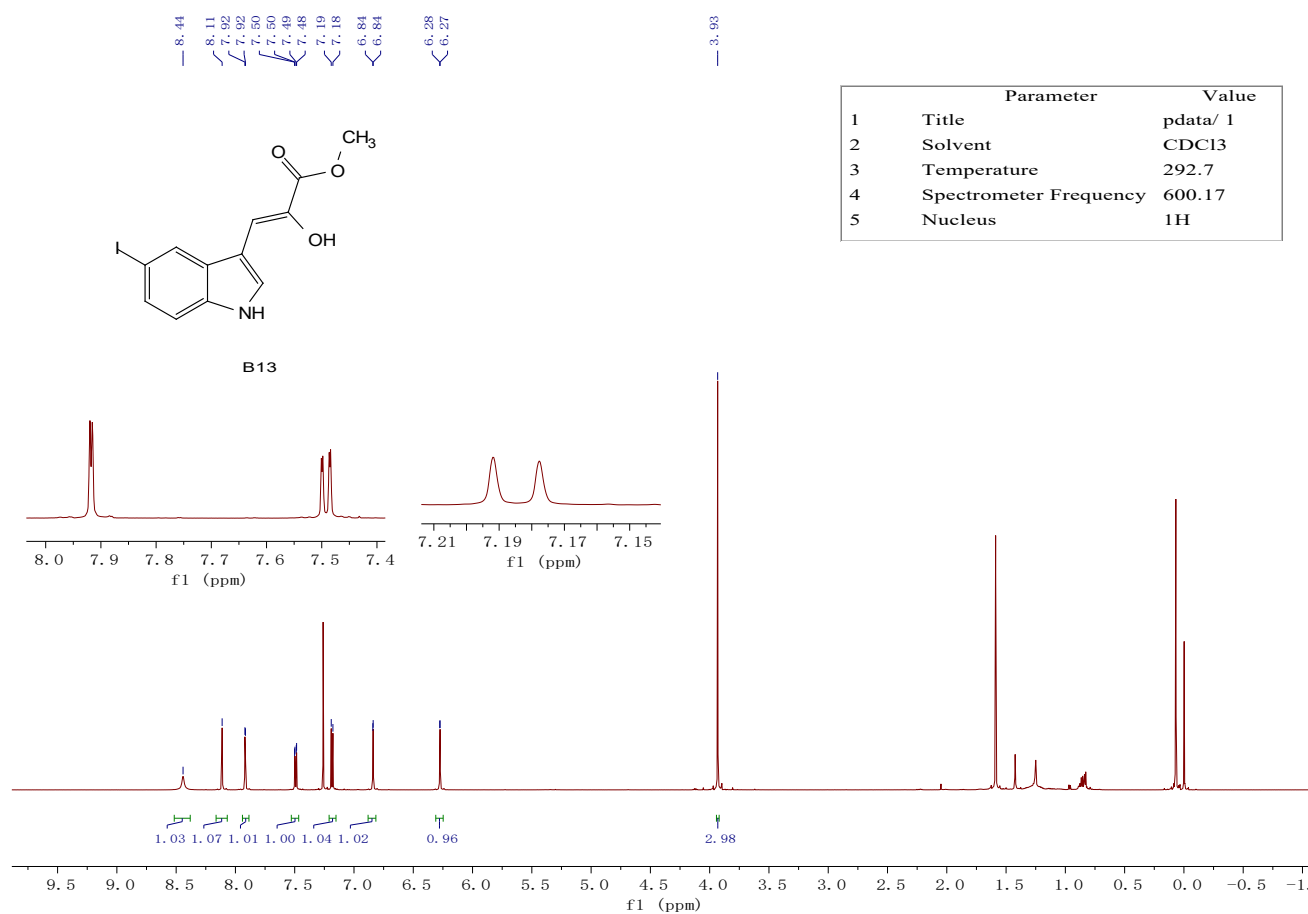


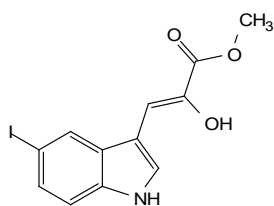


B12



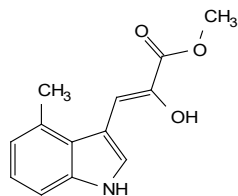
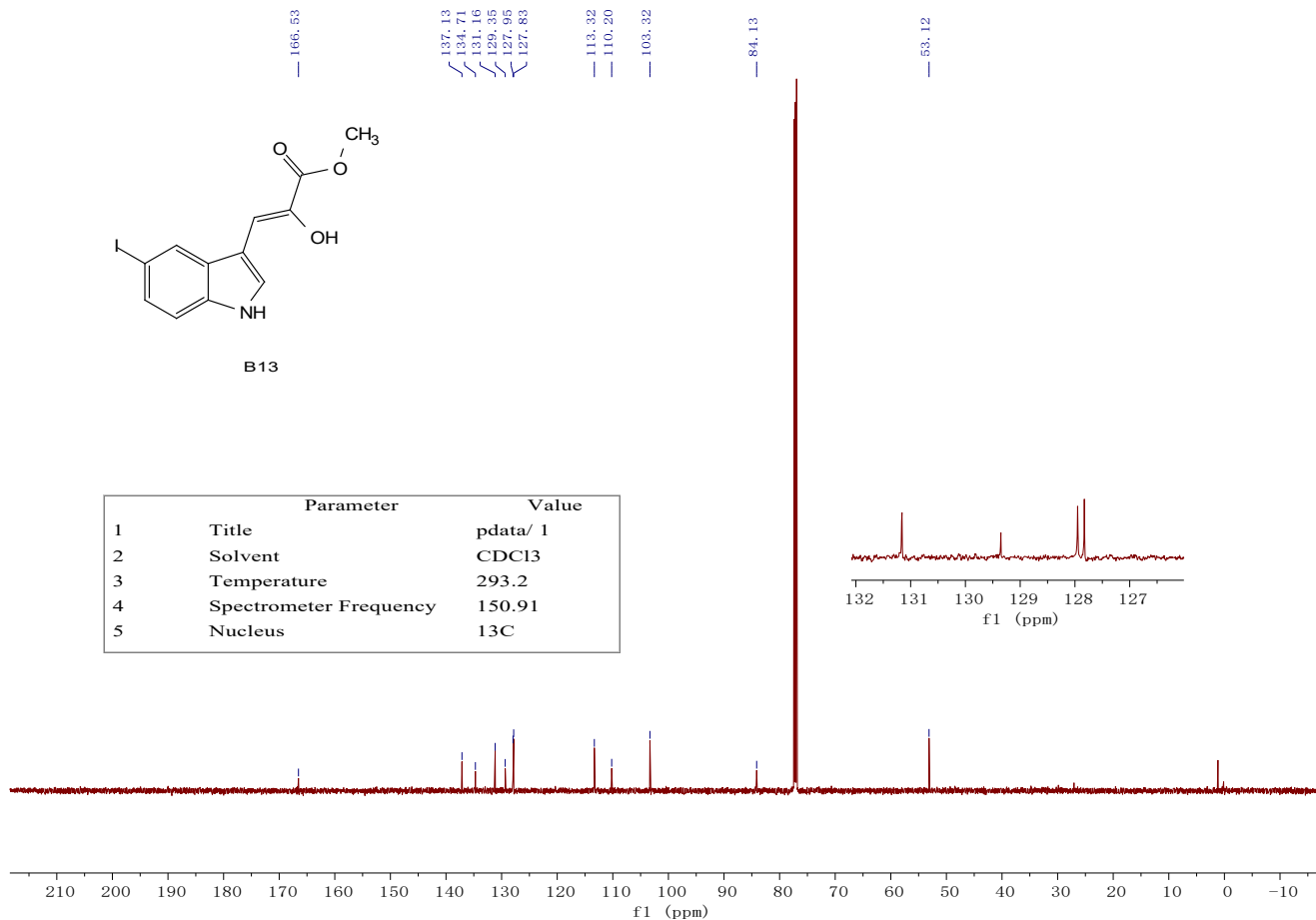
B13



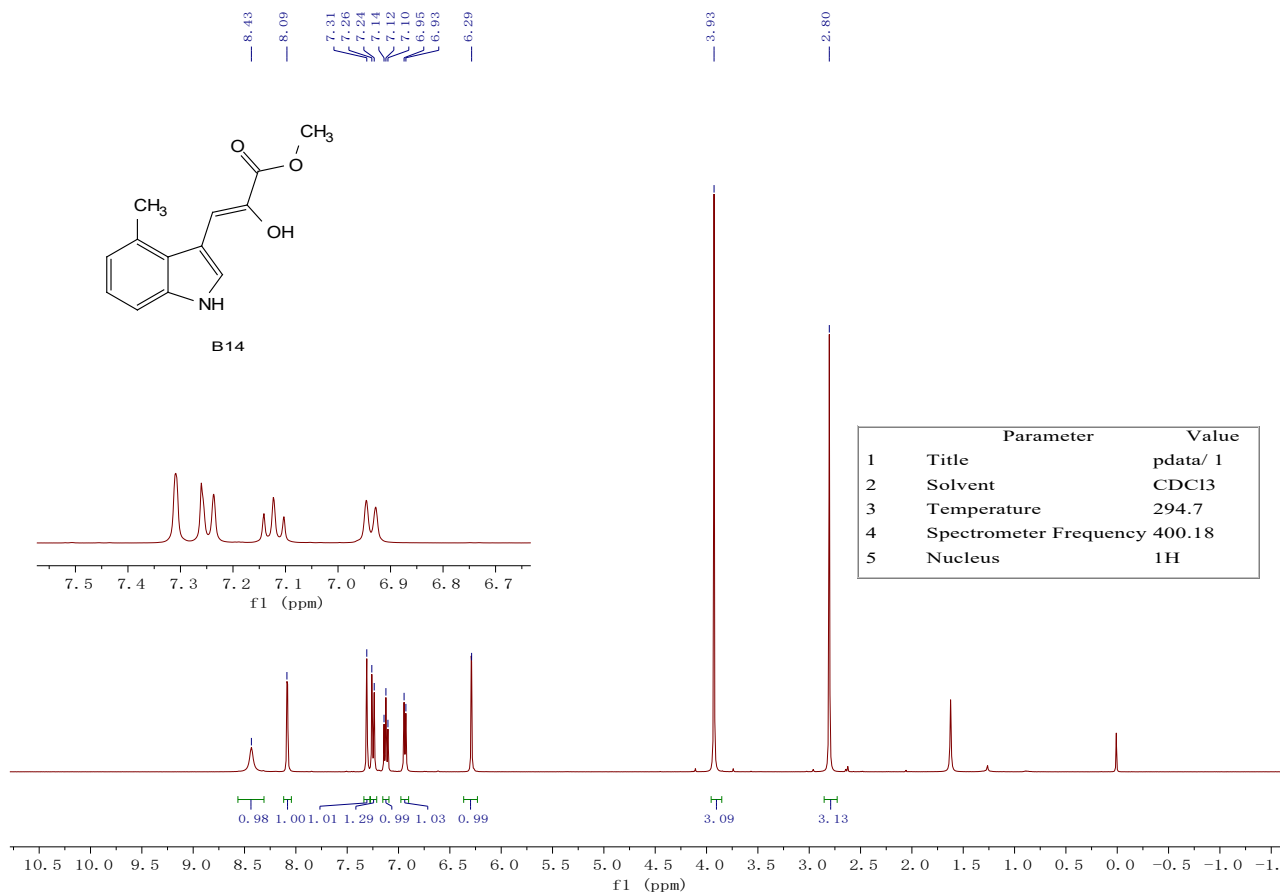


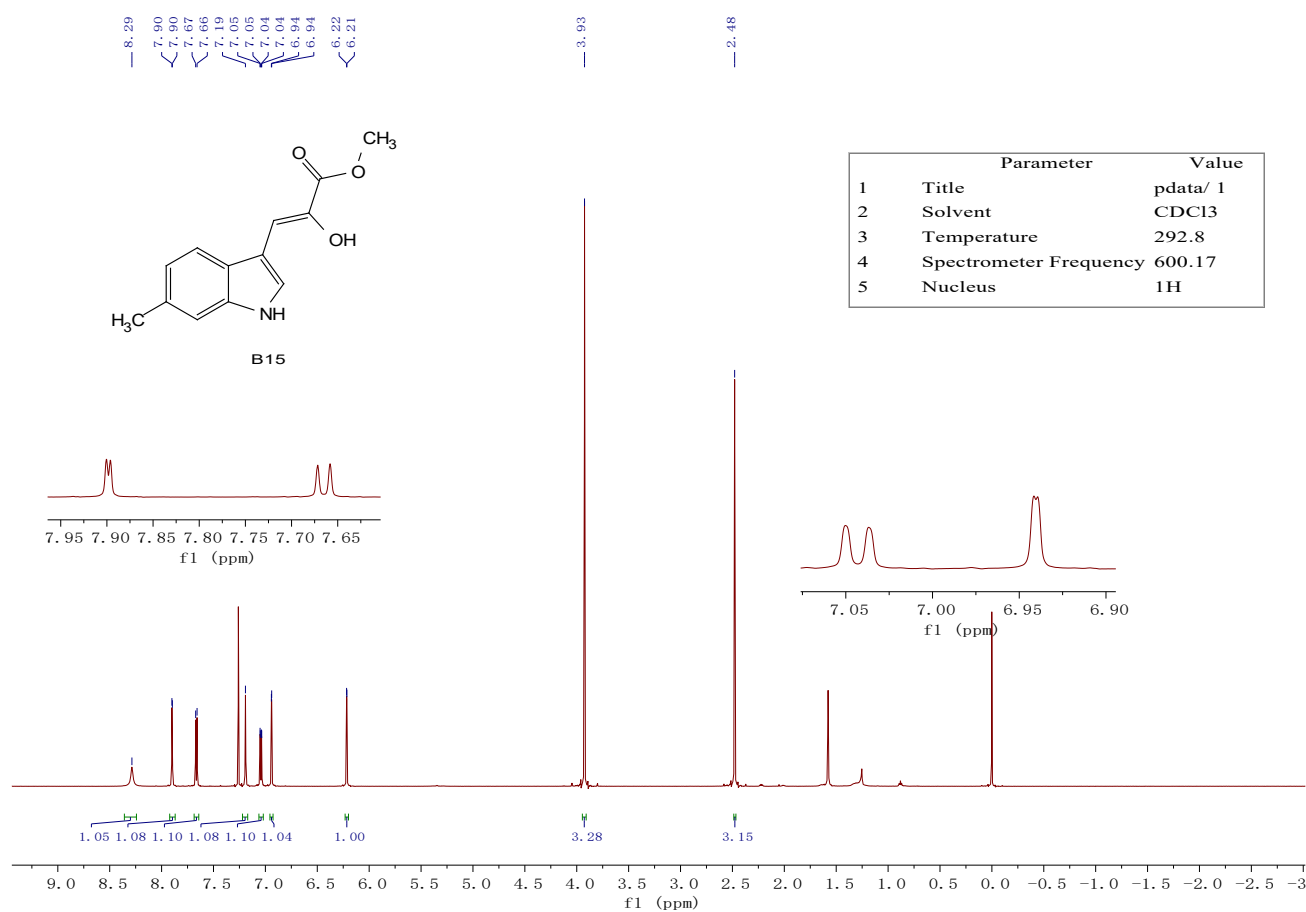
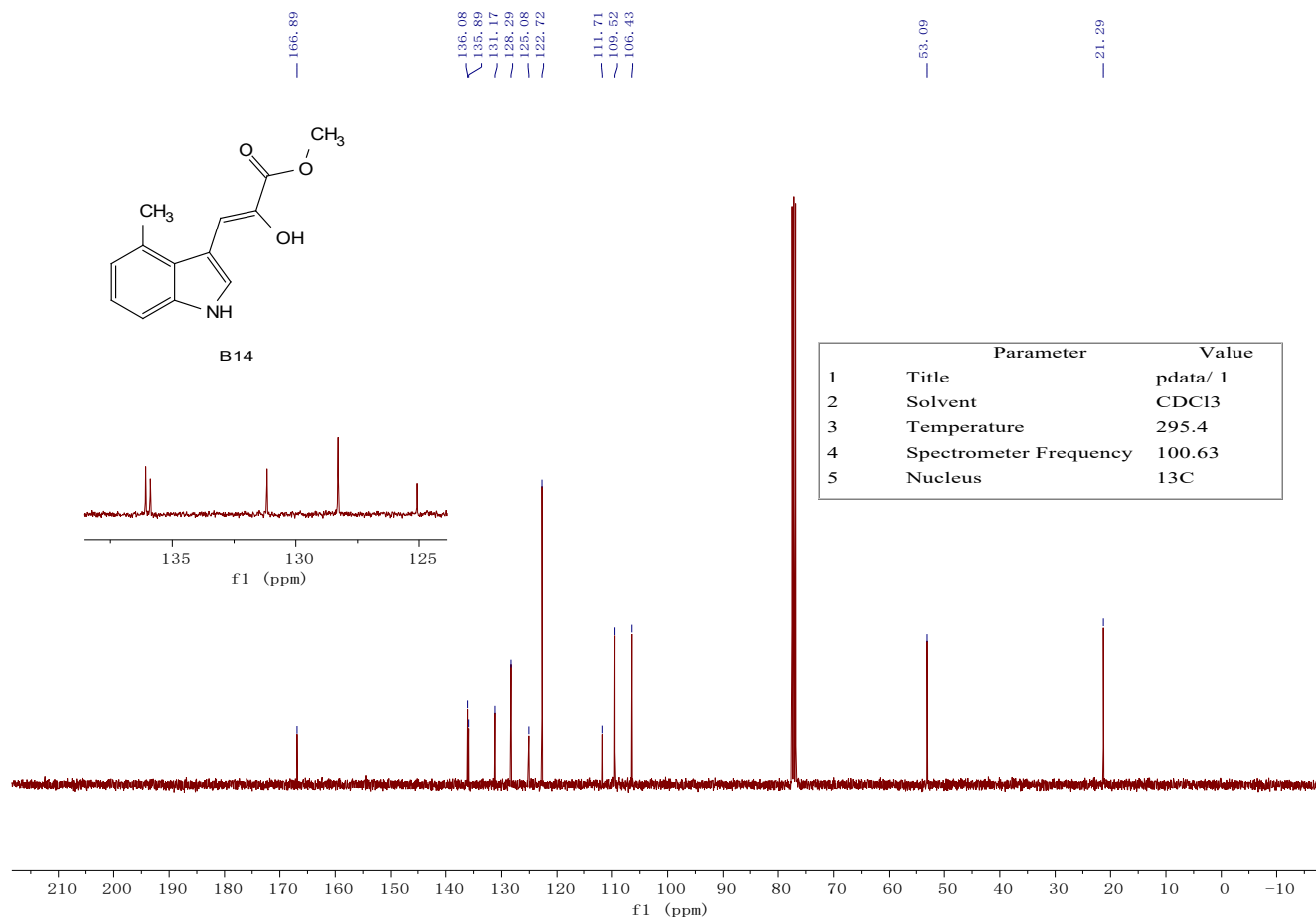
B13

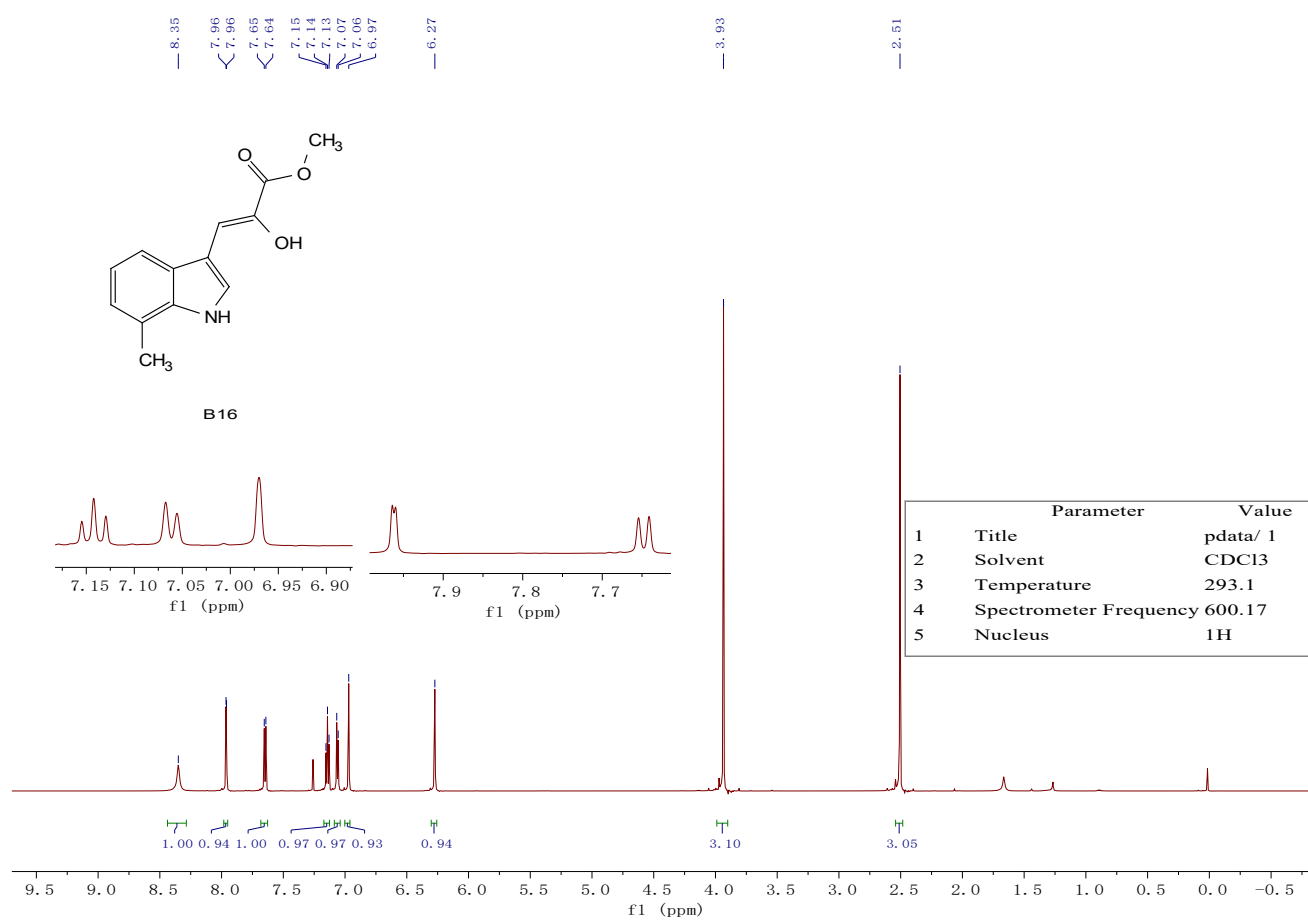
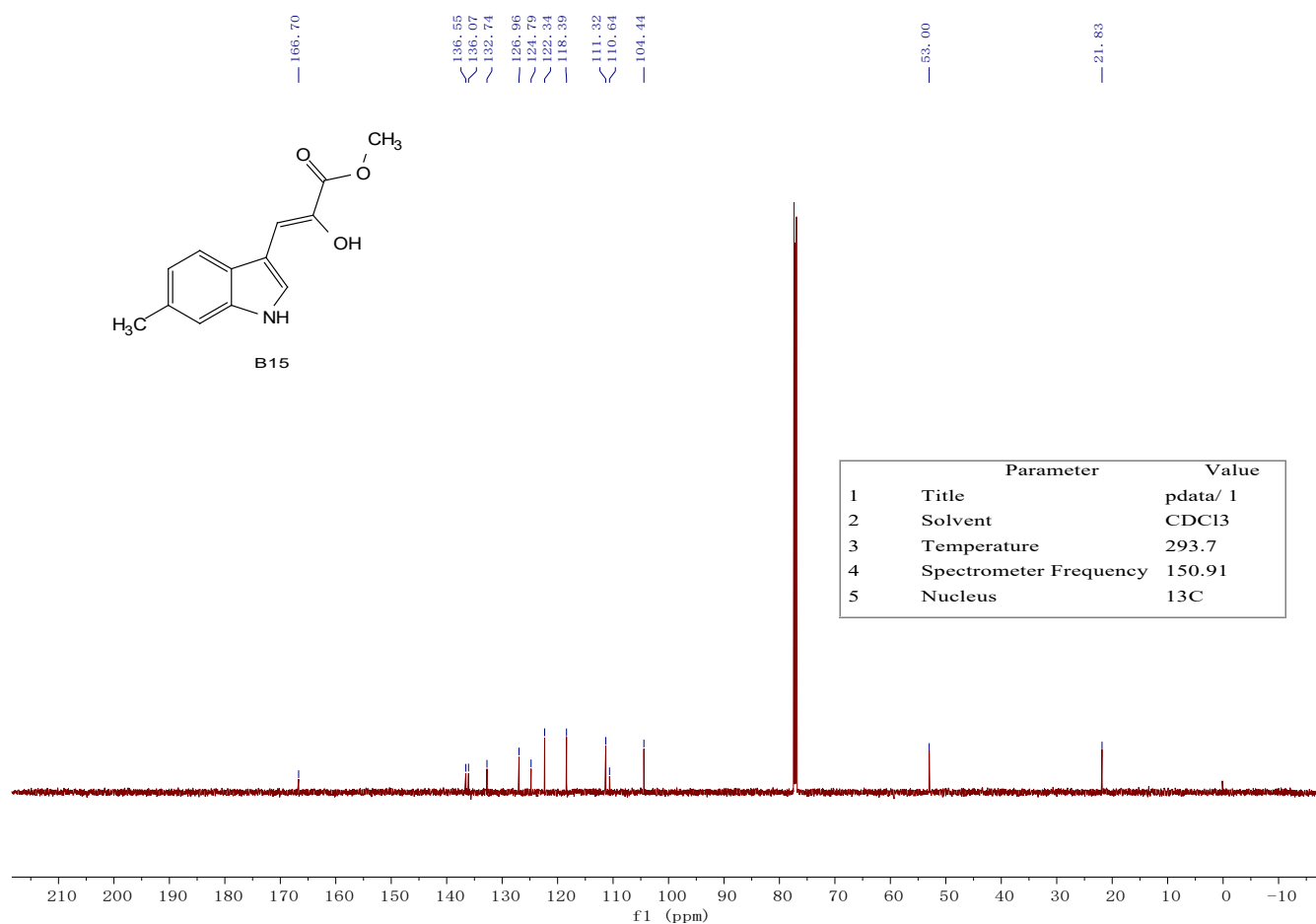
|   | Parameter              | Value    |
|---|------------------------|----------|
| 1 | Title                  | pdata/ 1 |
| 2 | Solvent                | CDCl3    |
| 3 | Temperature            | 293.2    |
| 4 | Spectrometer Frequency | 150.91   |
| 5 | Nucleus                | 13C      |

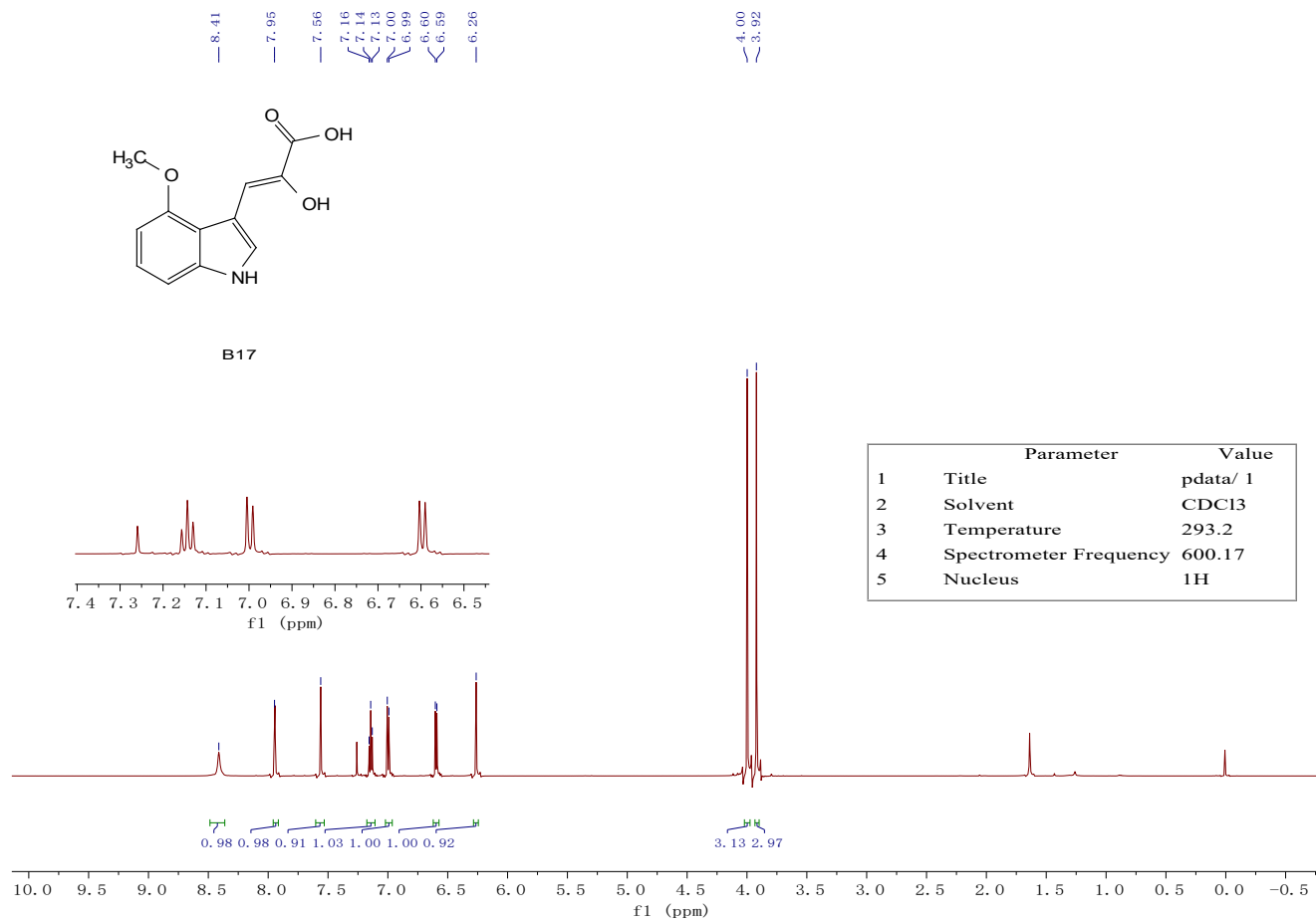
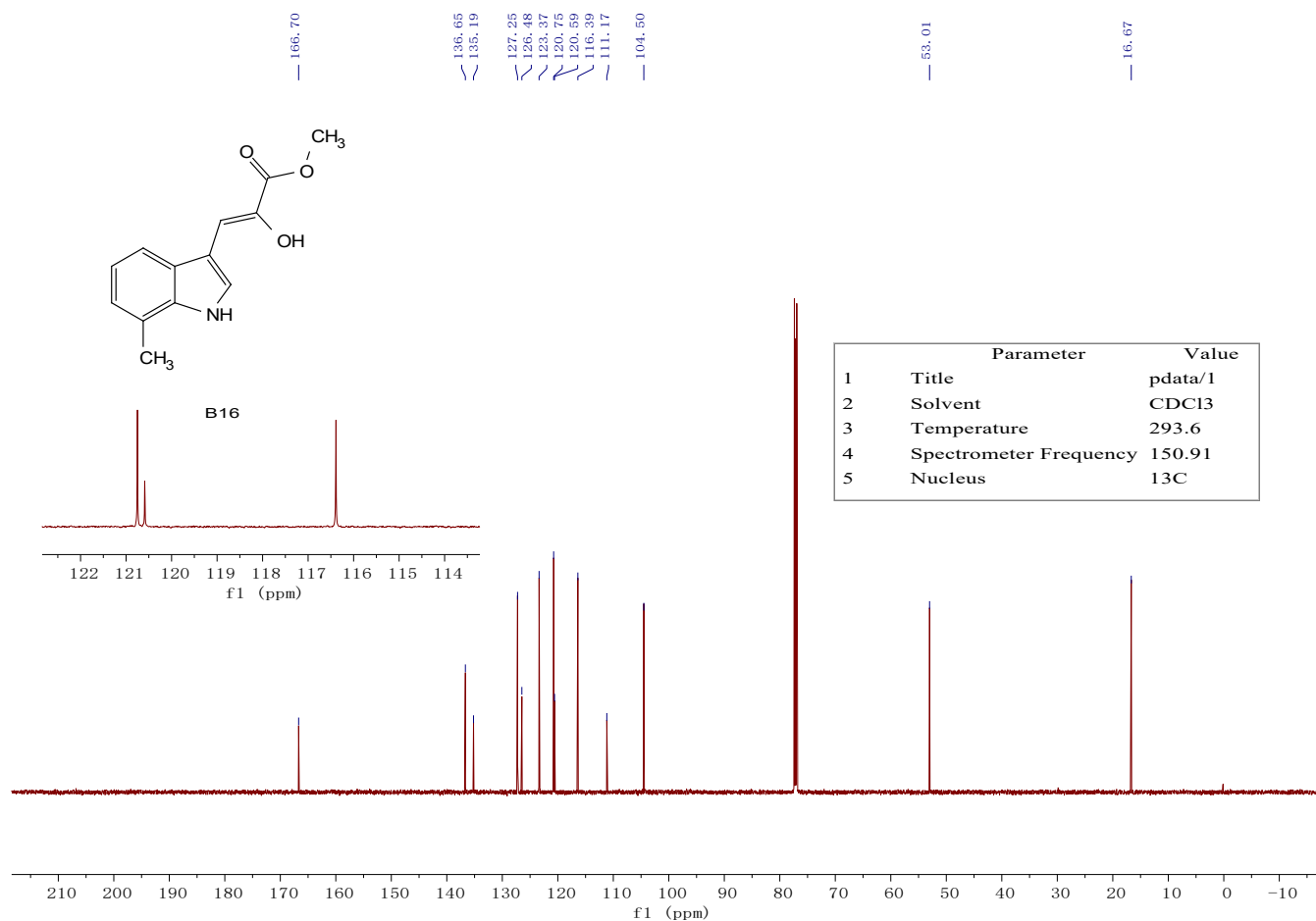


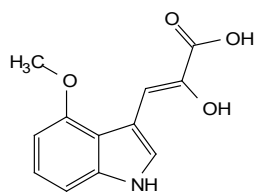
B14



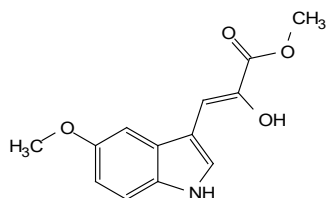
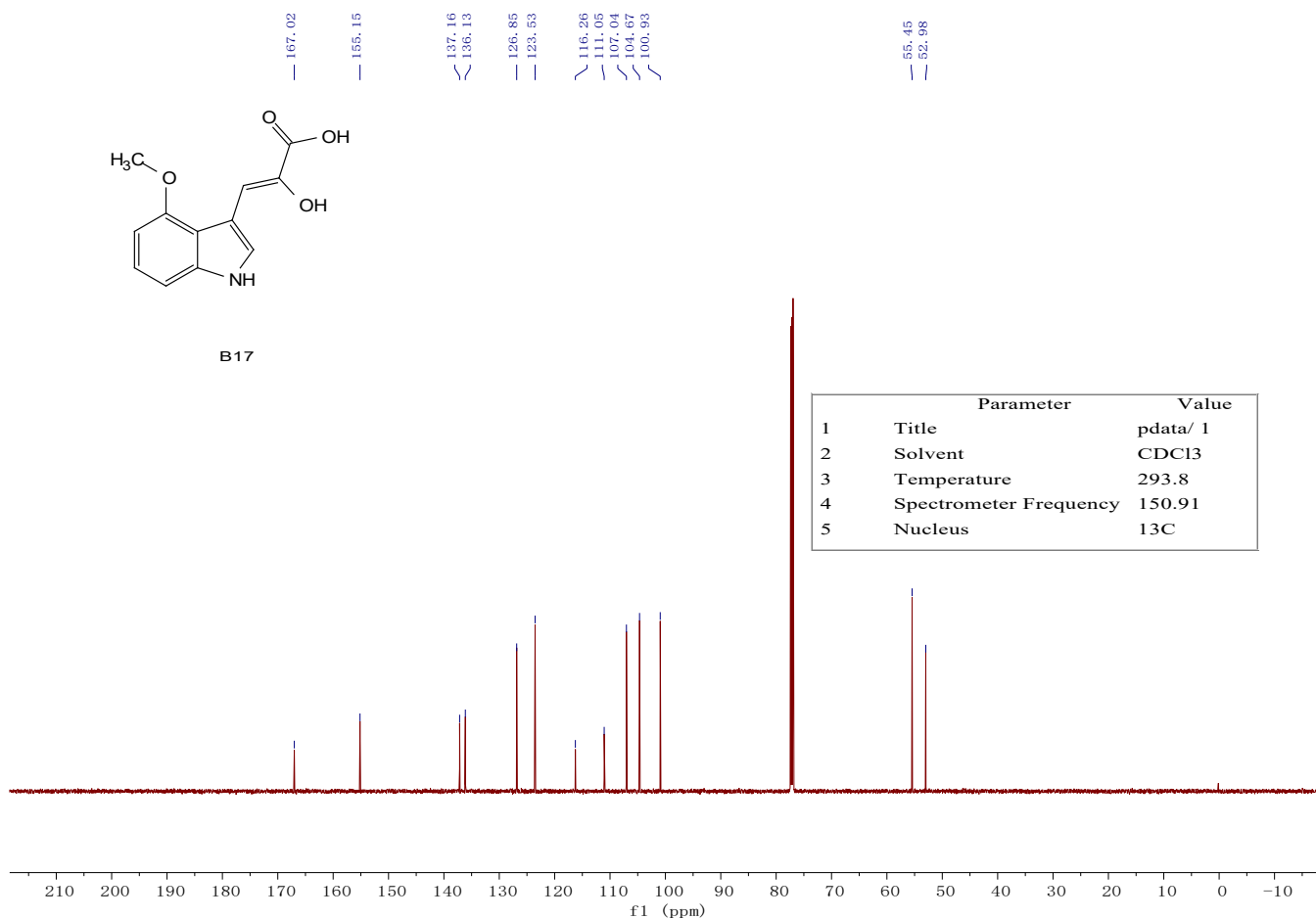




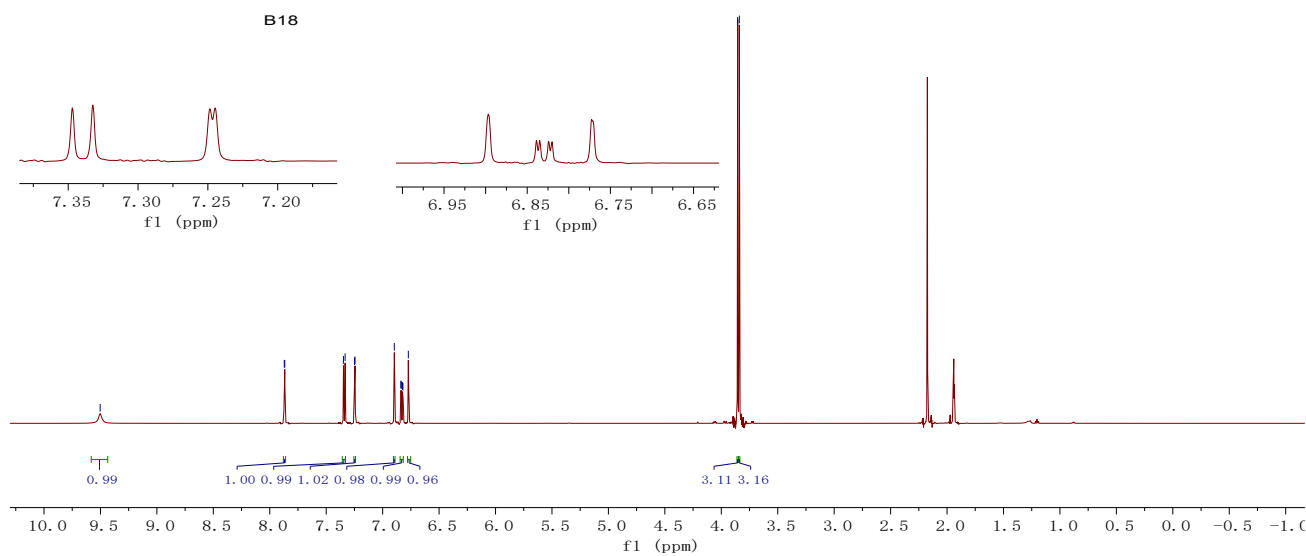


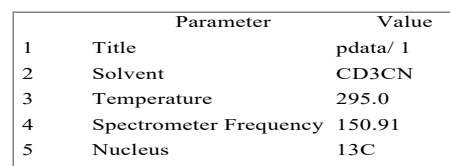


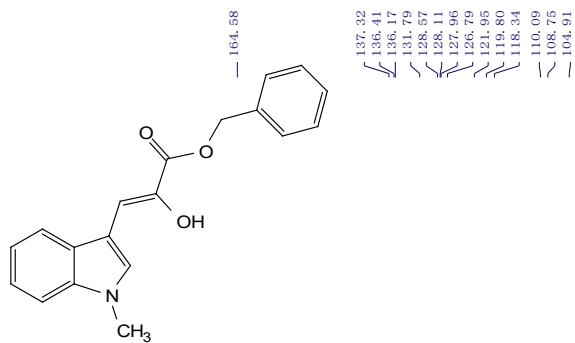
B17



B18

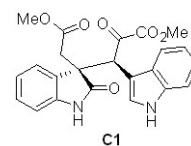
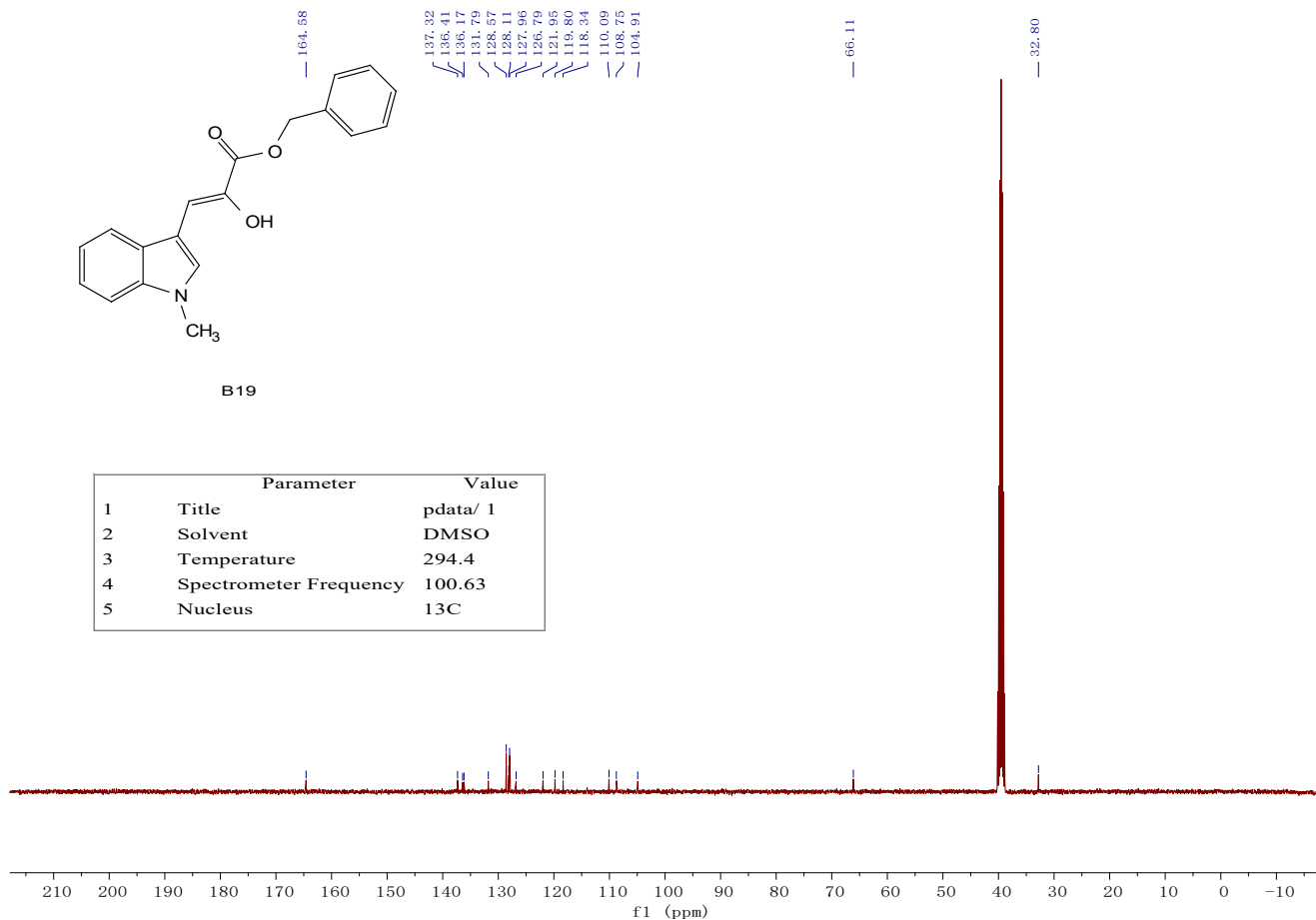




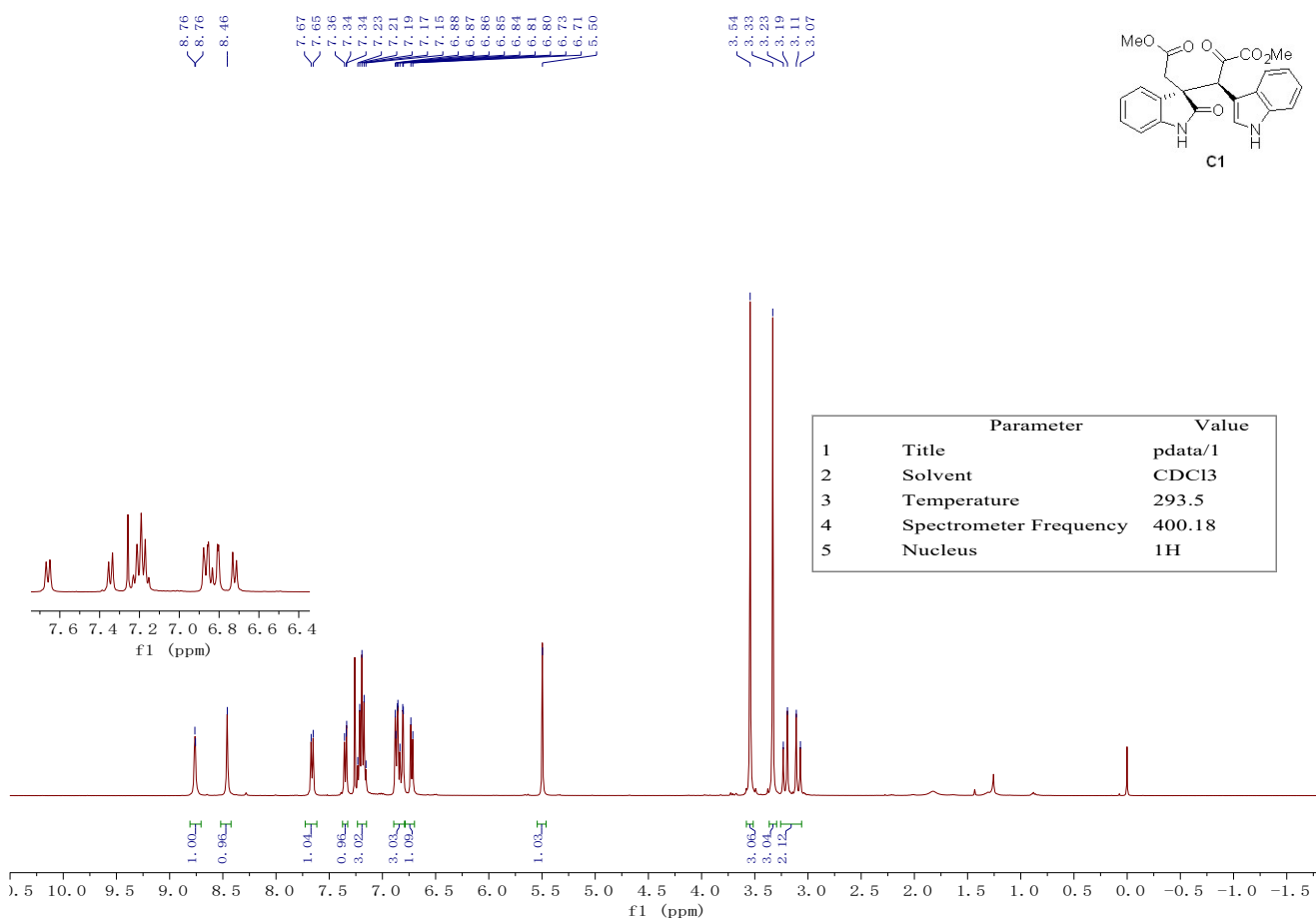


B19

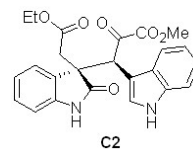
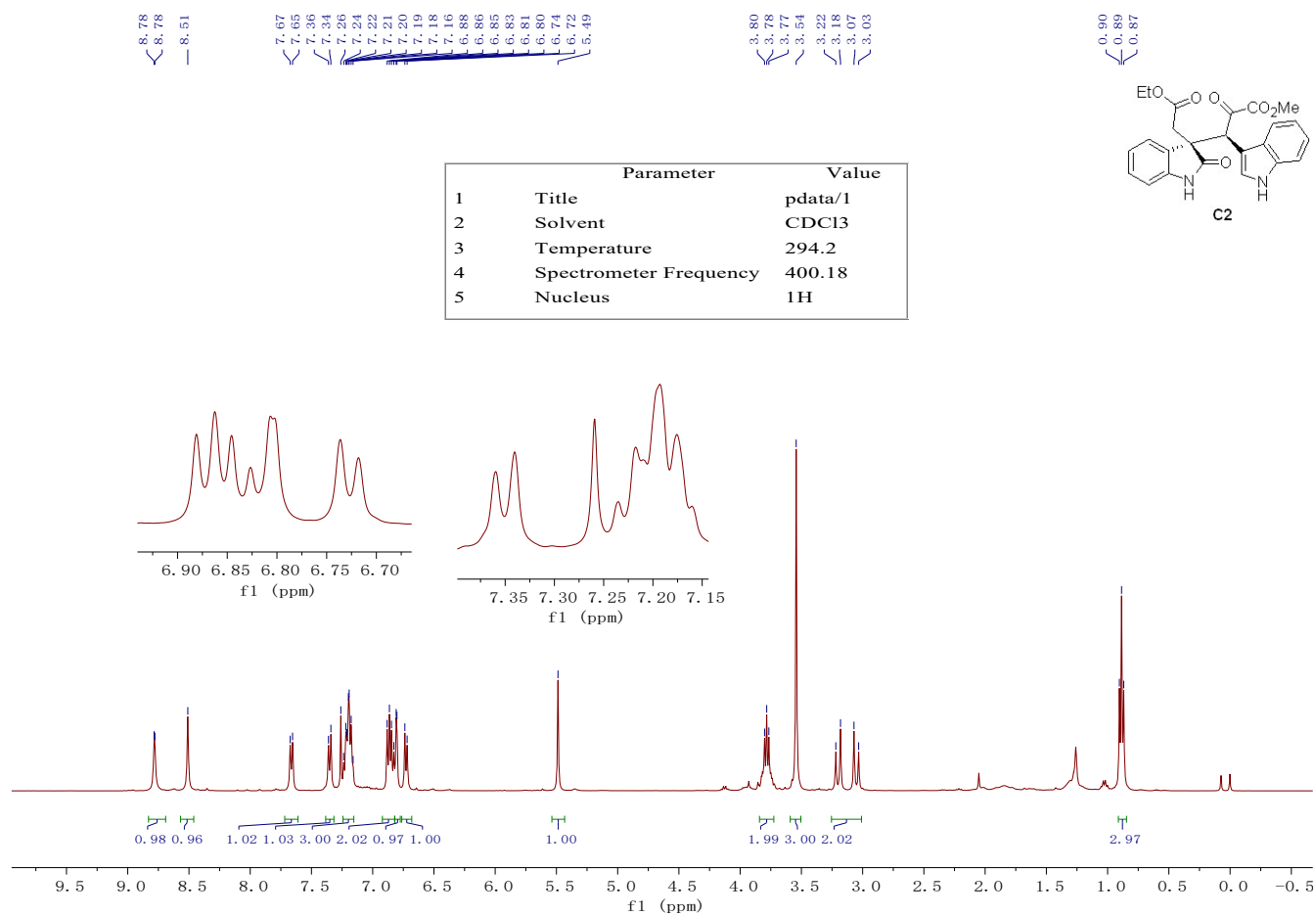
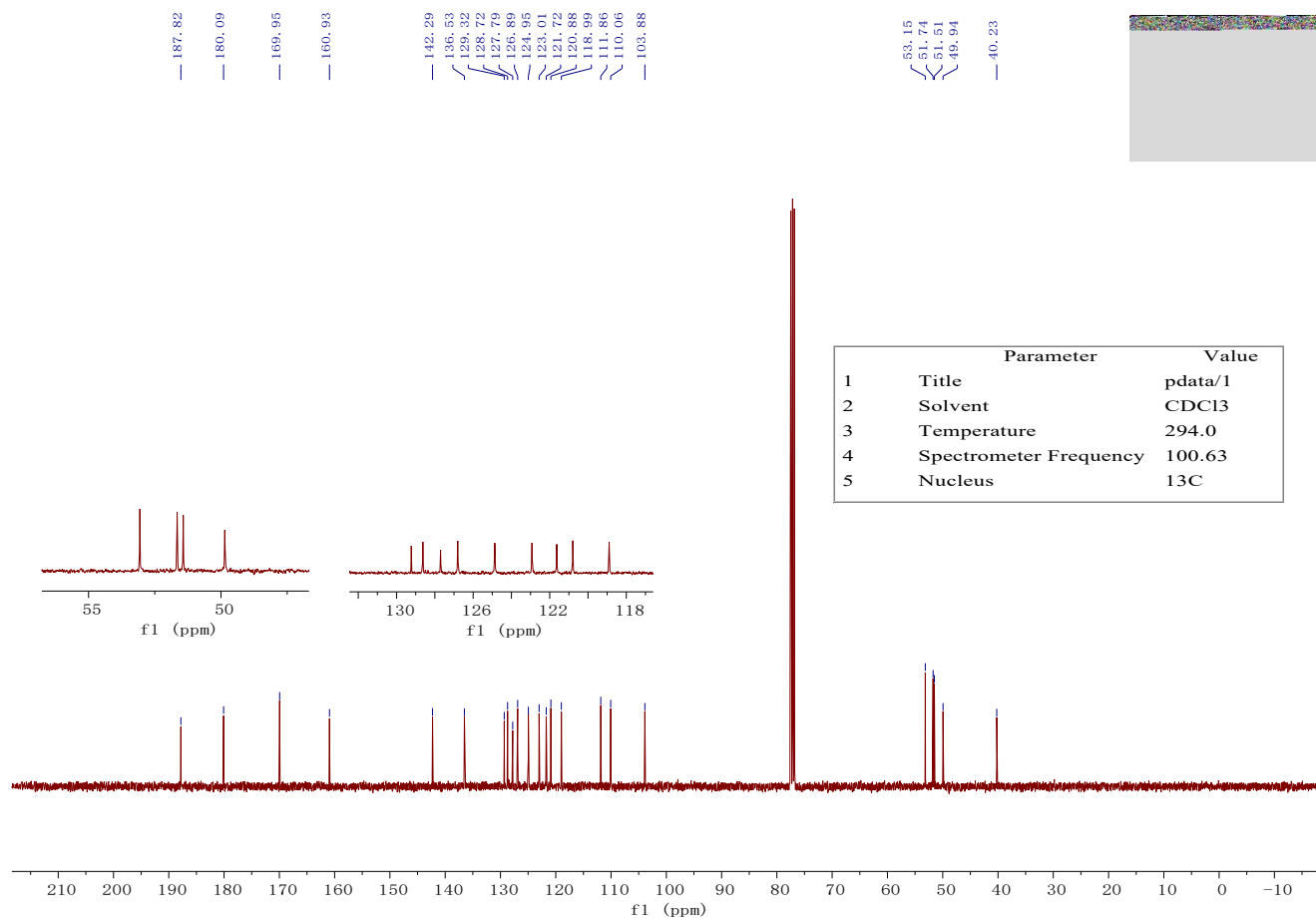
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|---|------------------------|-----------------|
| 1 | Title                  | pdata/ 1        |
| 2 | Solvent                | DMSO            |
| 3 | Temperature            | 294.4           |
| 4 | Spectrometer Frequency | 100.63          |
| 5 | Nucleus                | <sup>13</sup> C |

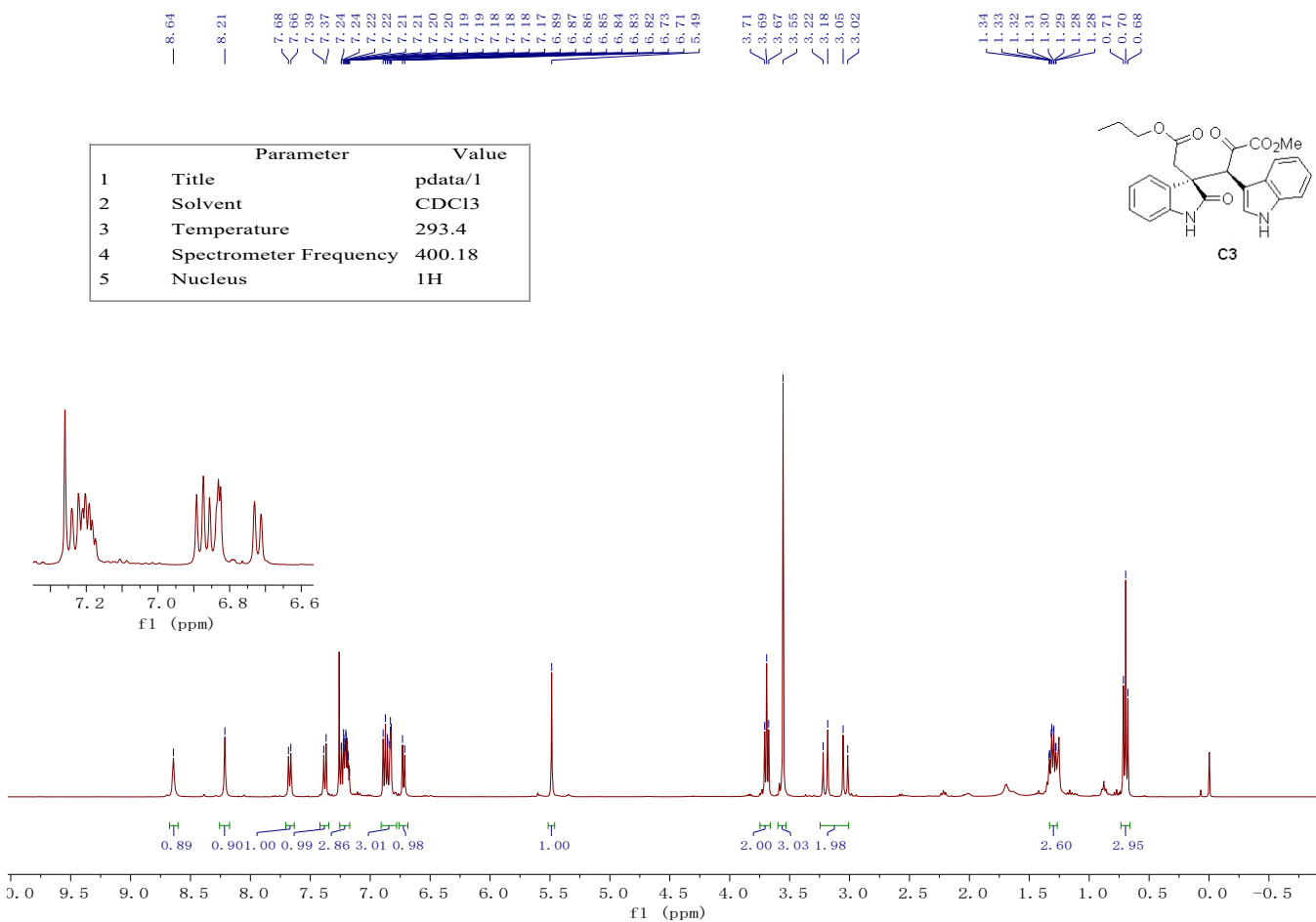
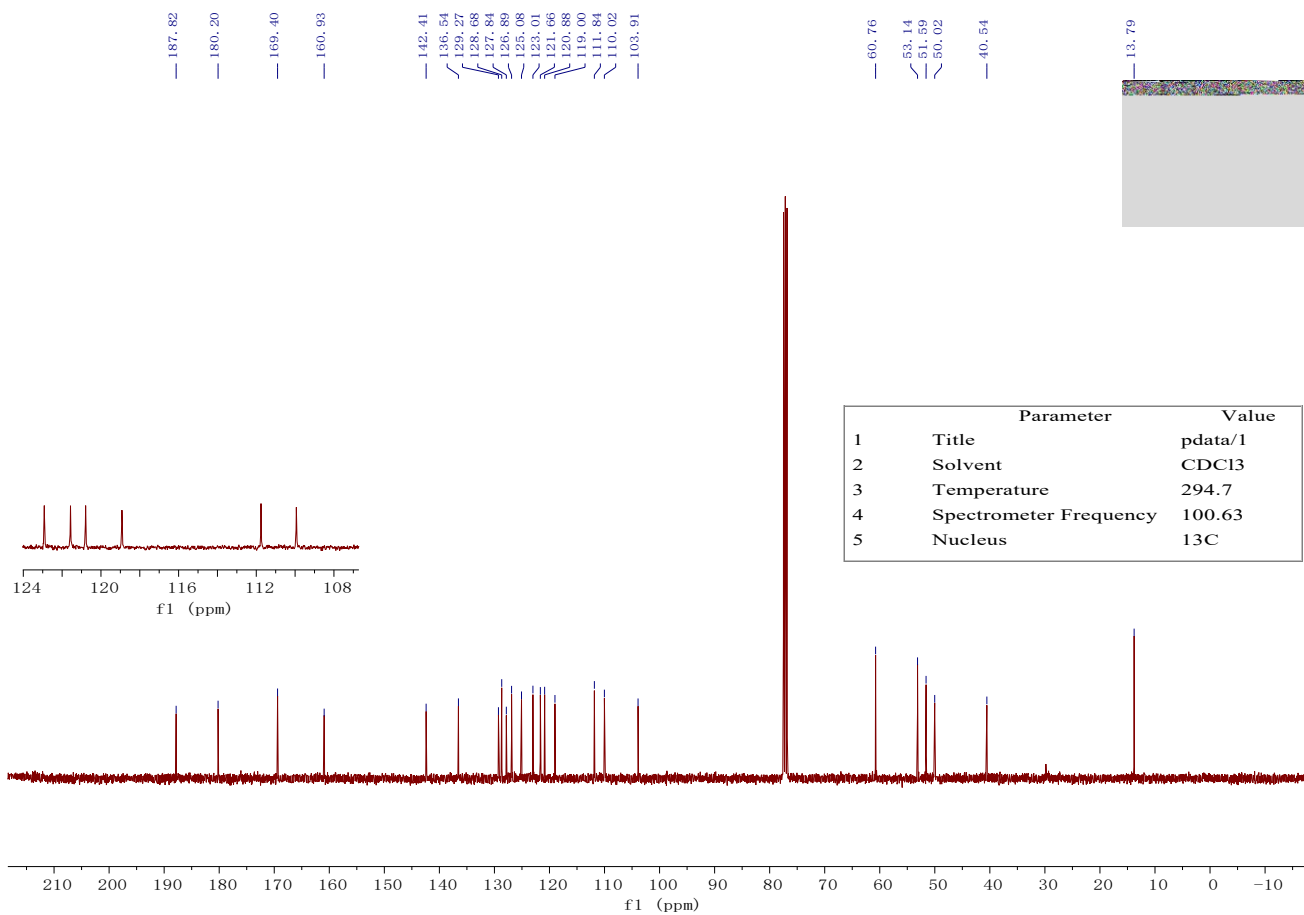


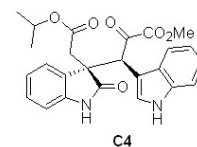
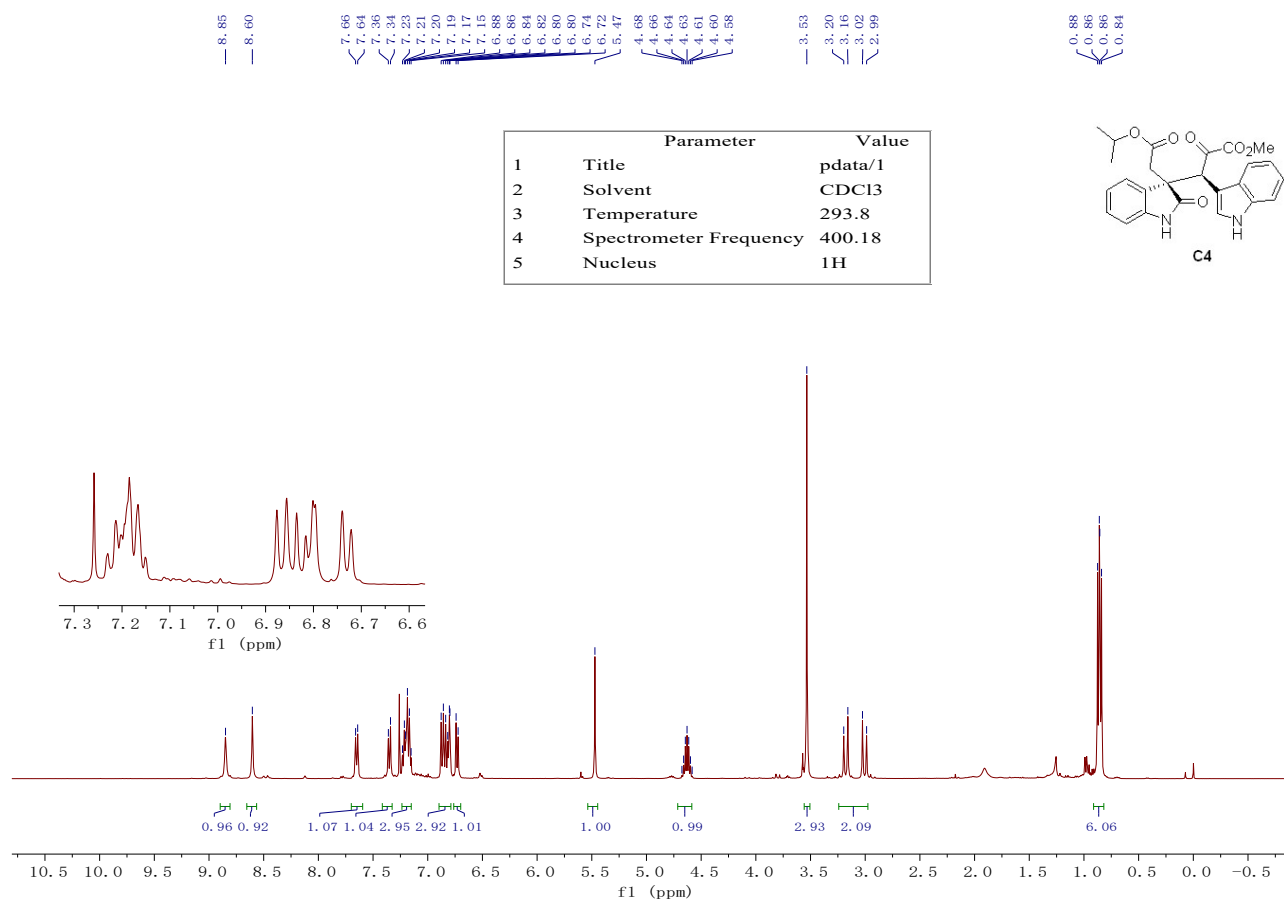
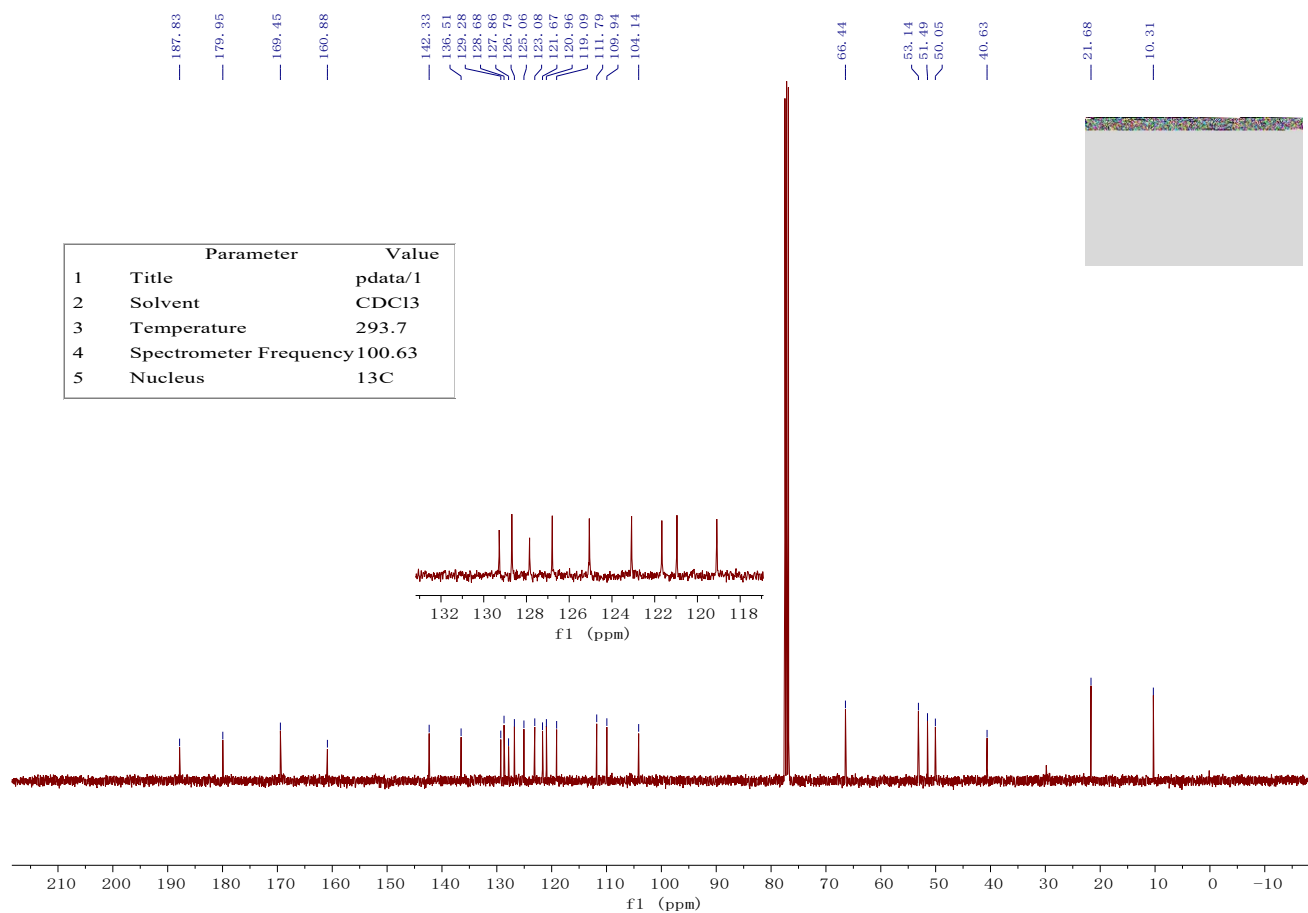
C1

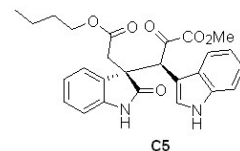
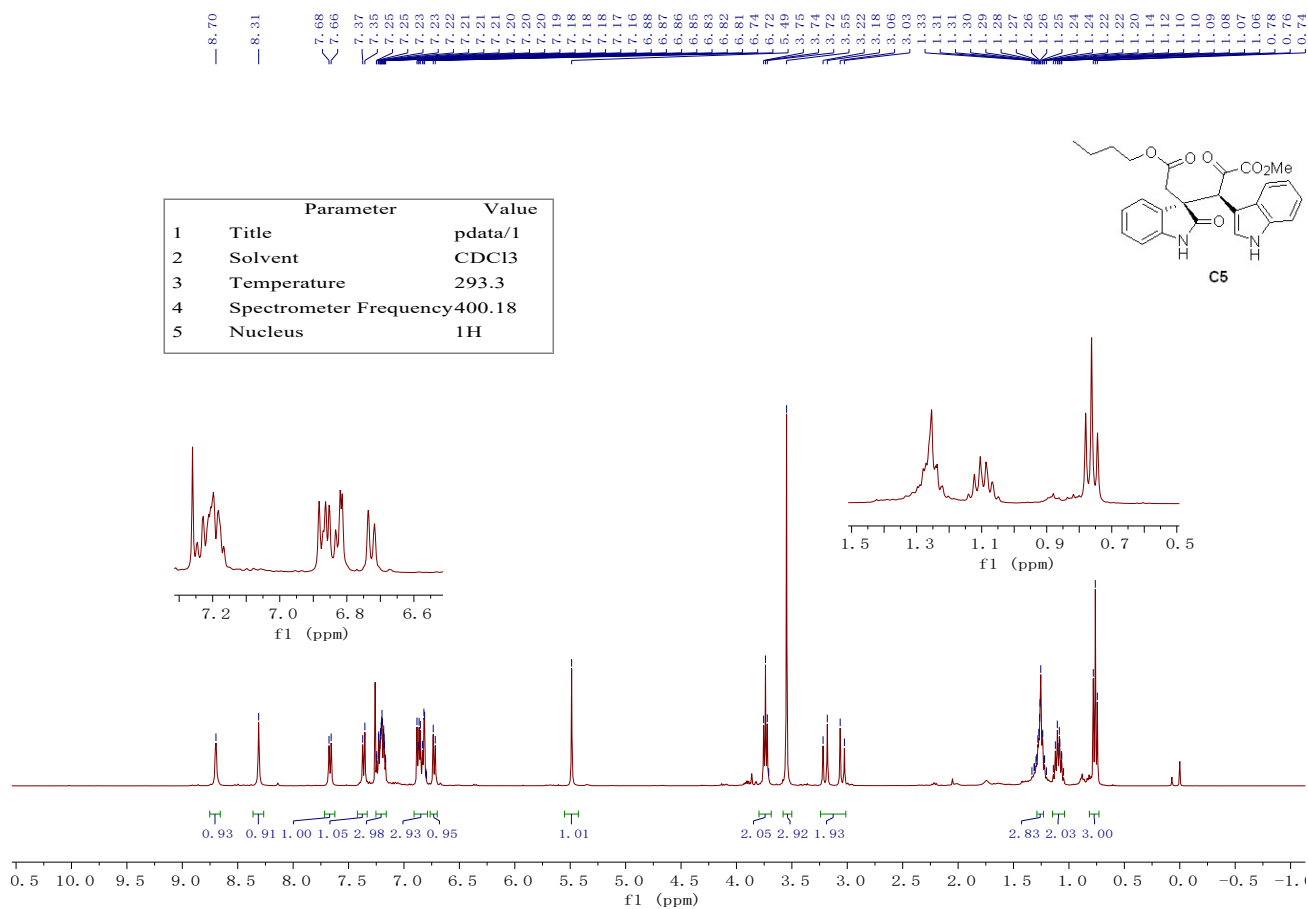
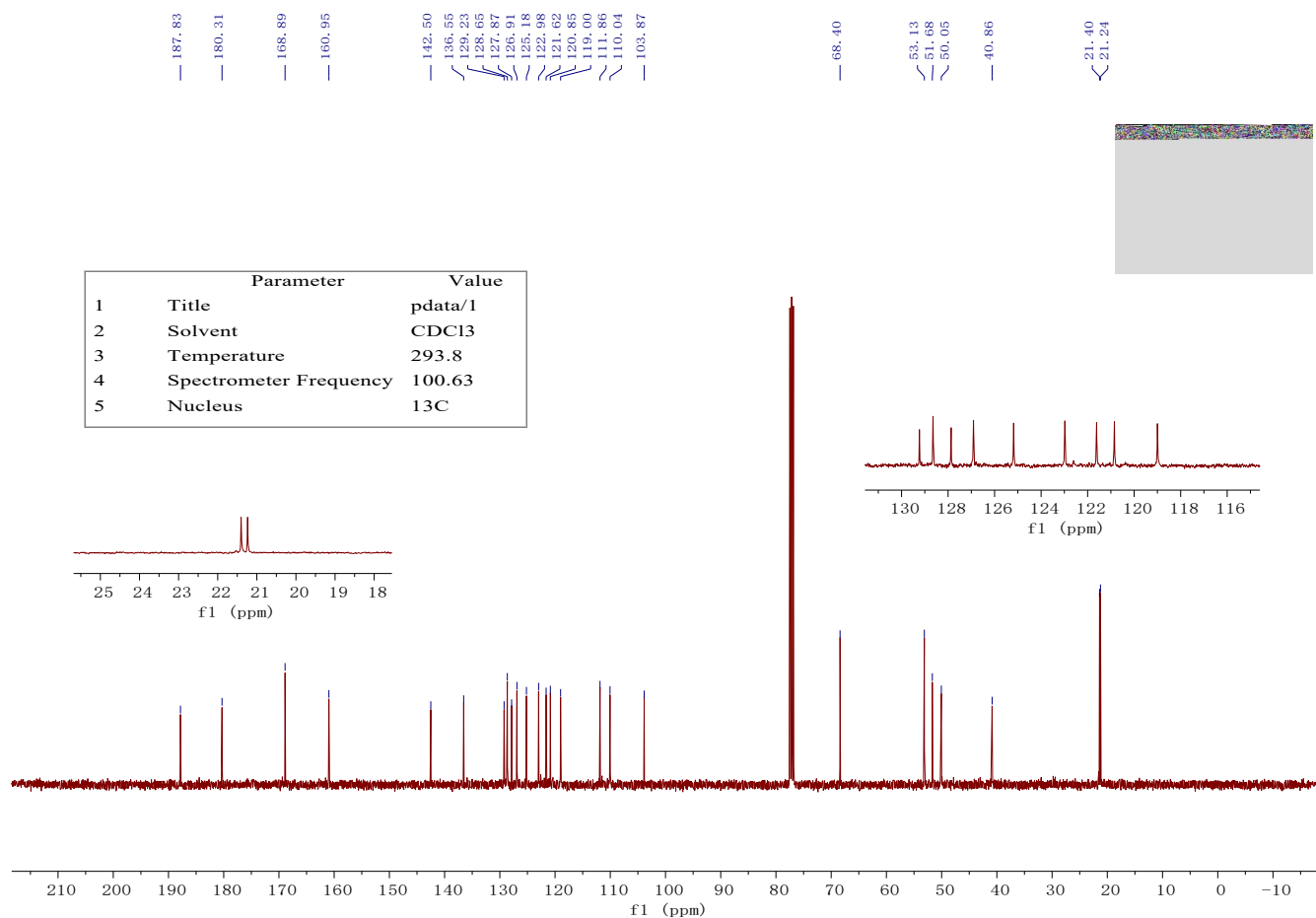


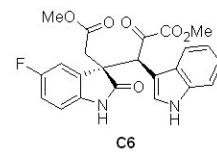
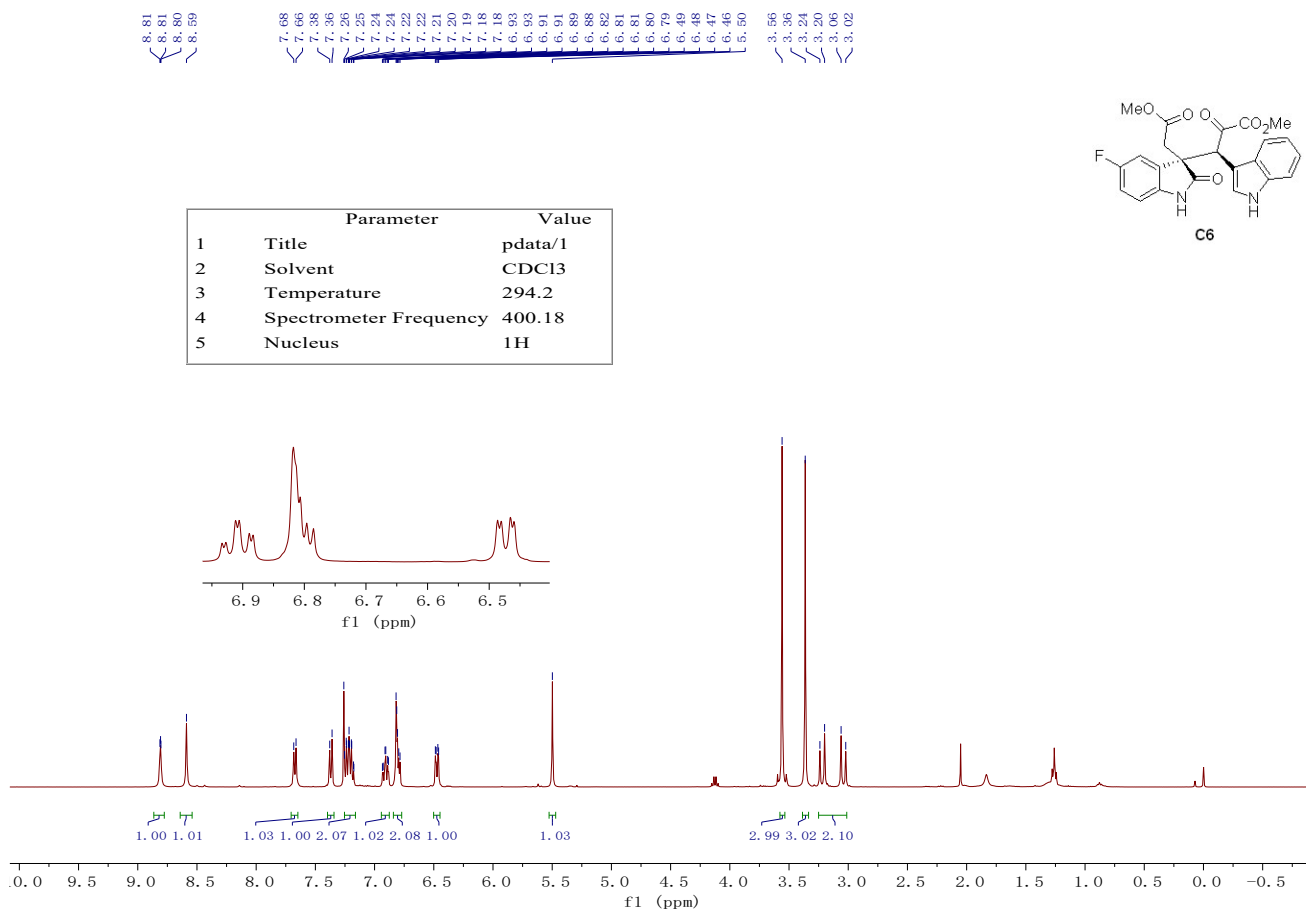
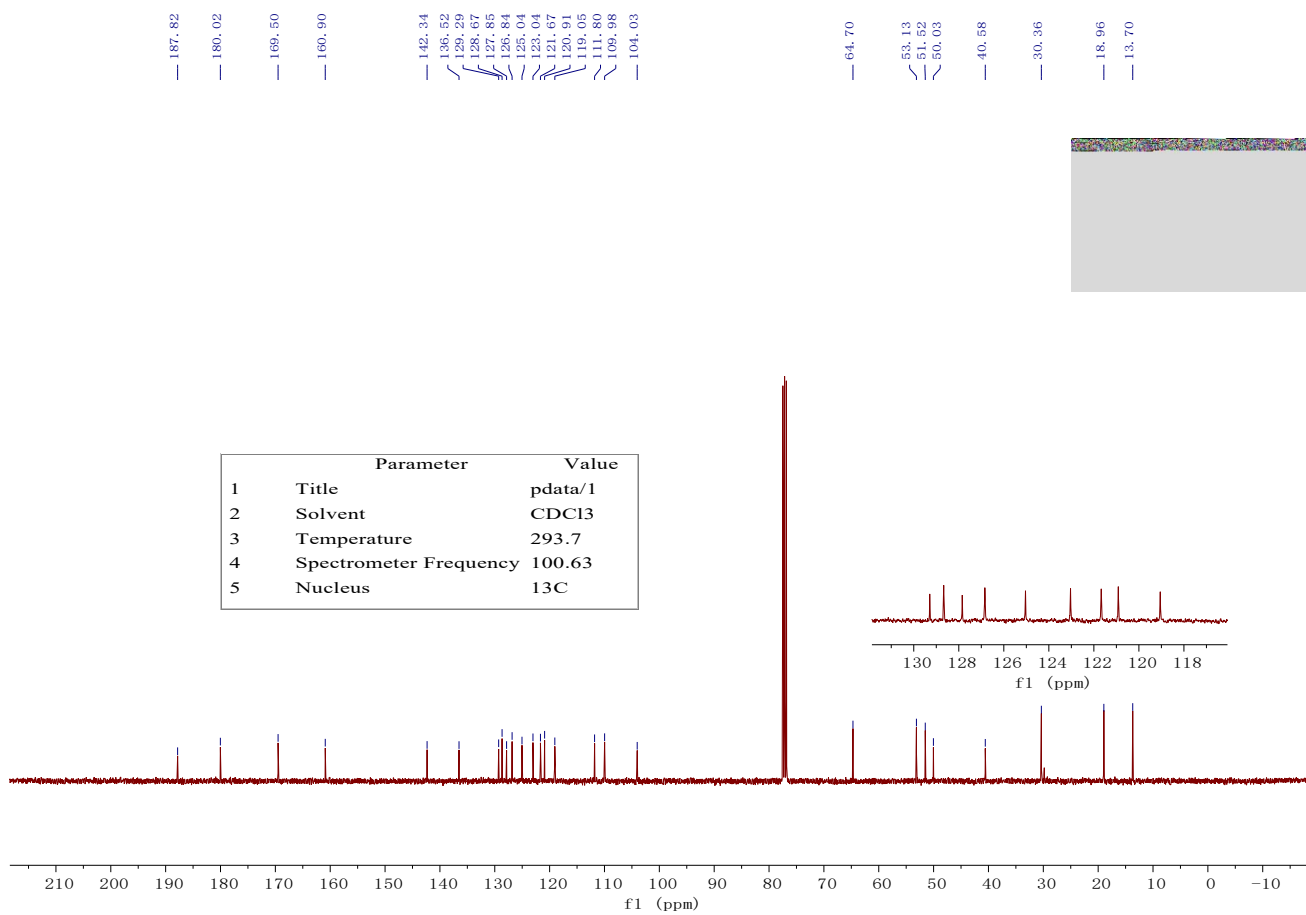
|   | Parameter              | Value          |
|---|------------------------|----------------|
| 1 | Title                  | pdata/1        |
| 2 | Solvent                | CDCl3          |
| 3 | Temperature            | 293.5          |
| 4 | Spectrometer Frequency | 400.18         |
| 5 | Nucleus                | <sup>1</sup> H |

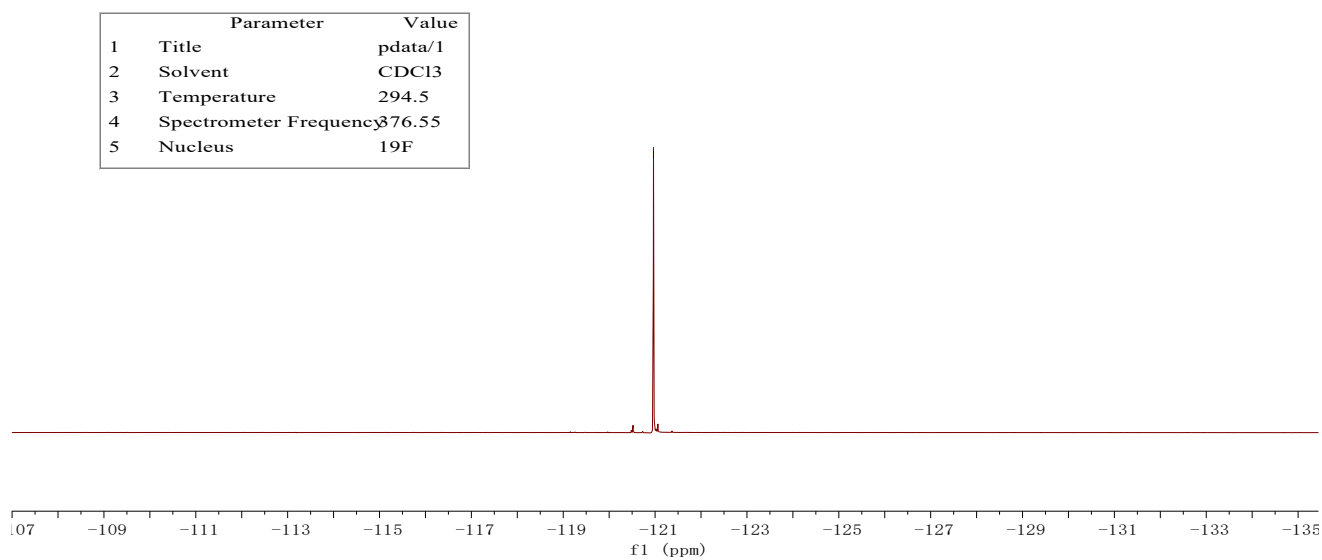
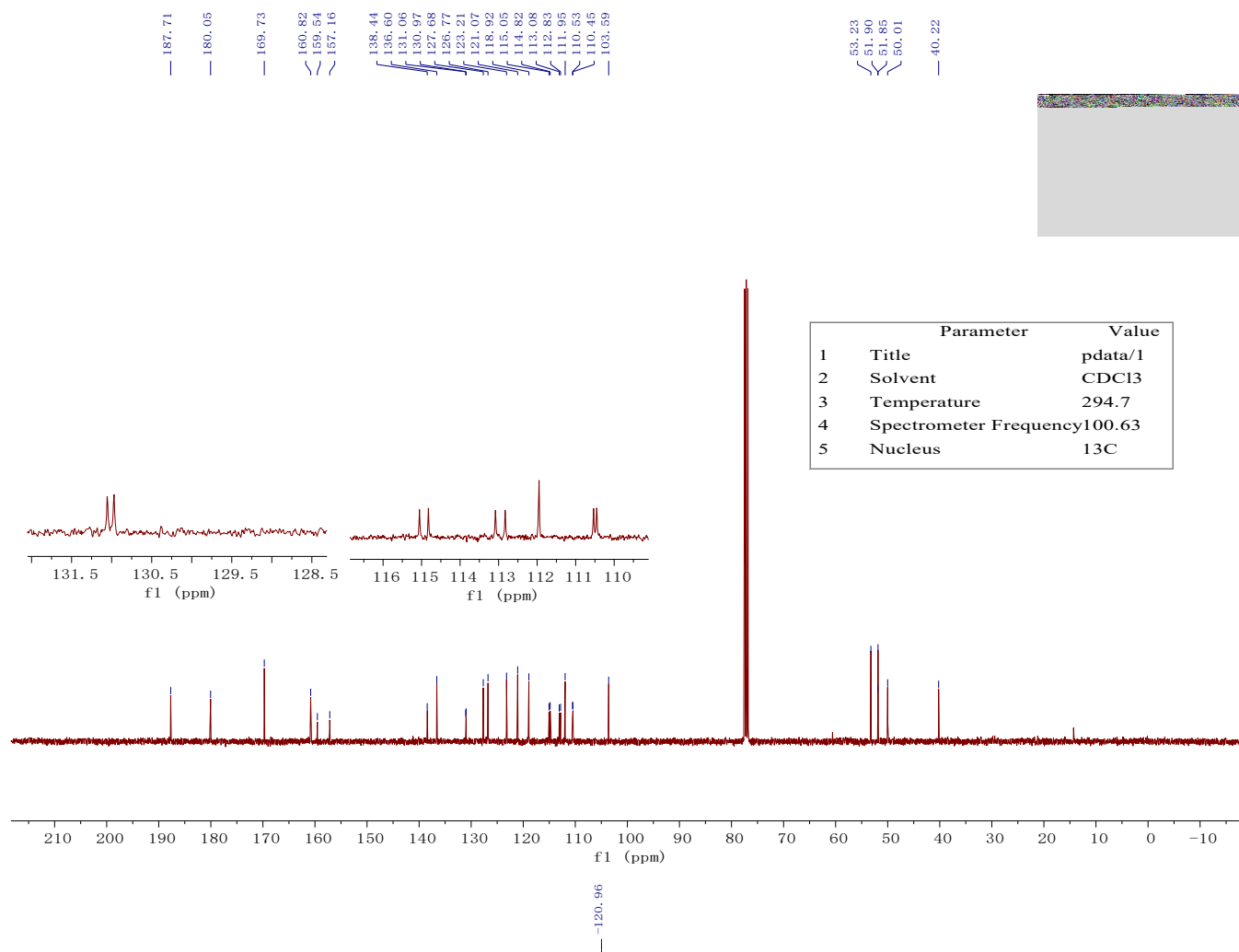


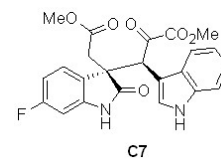




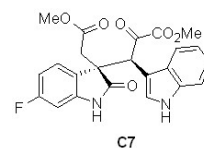
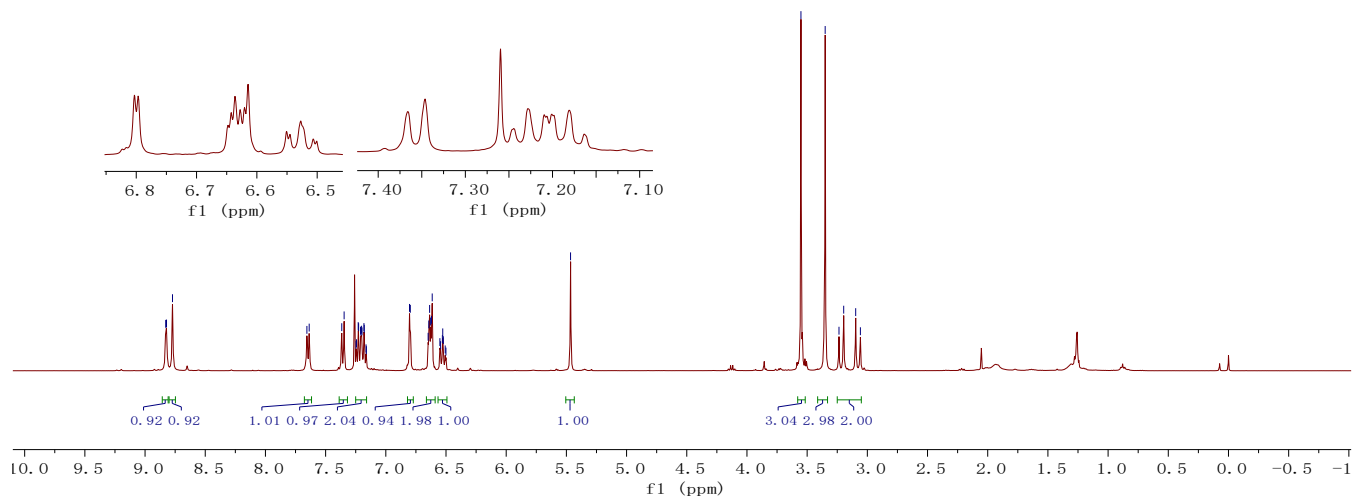




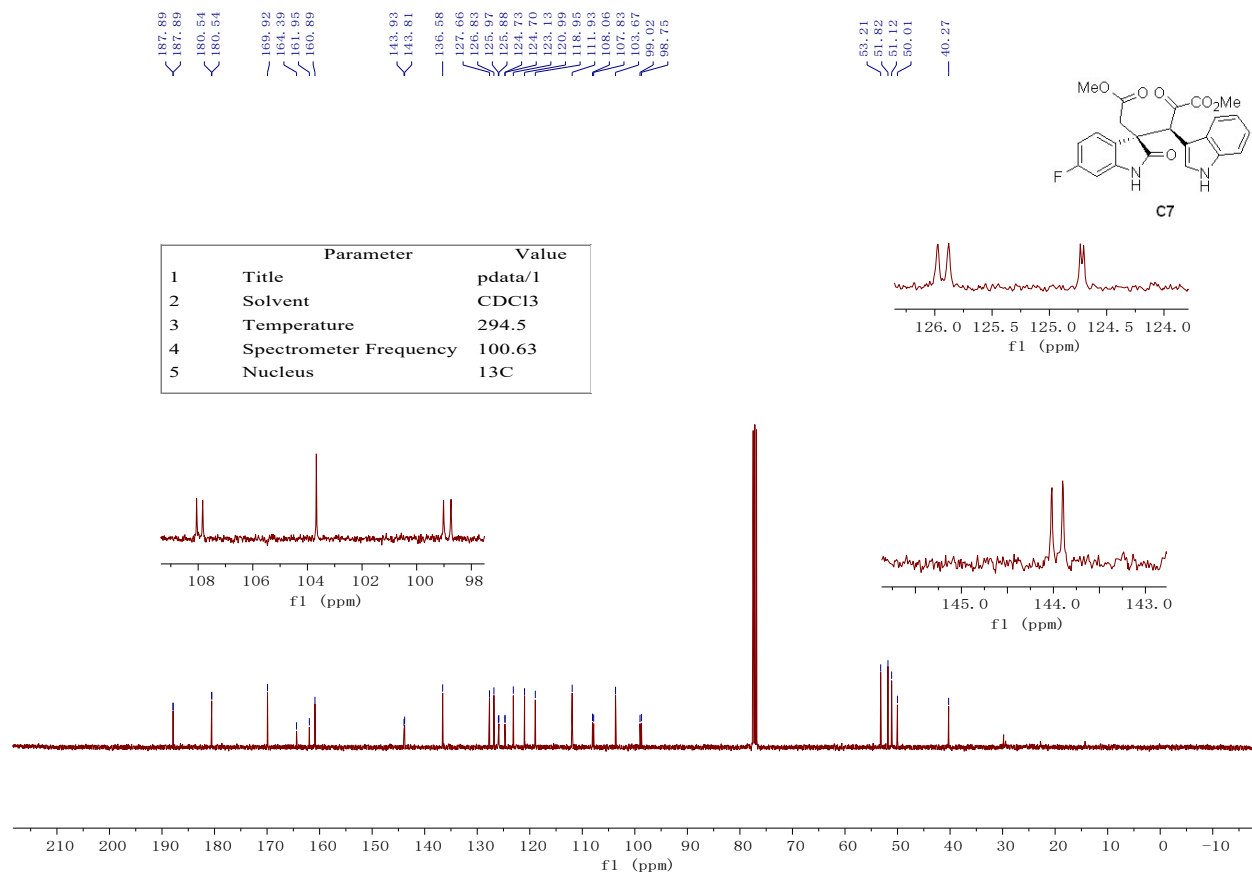


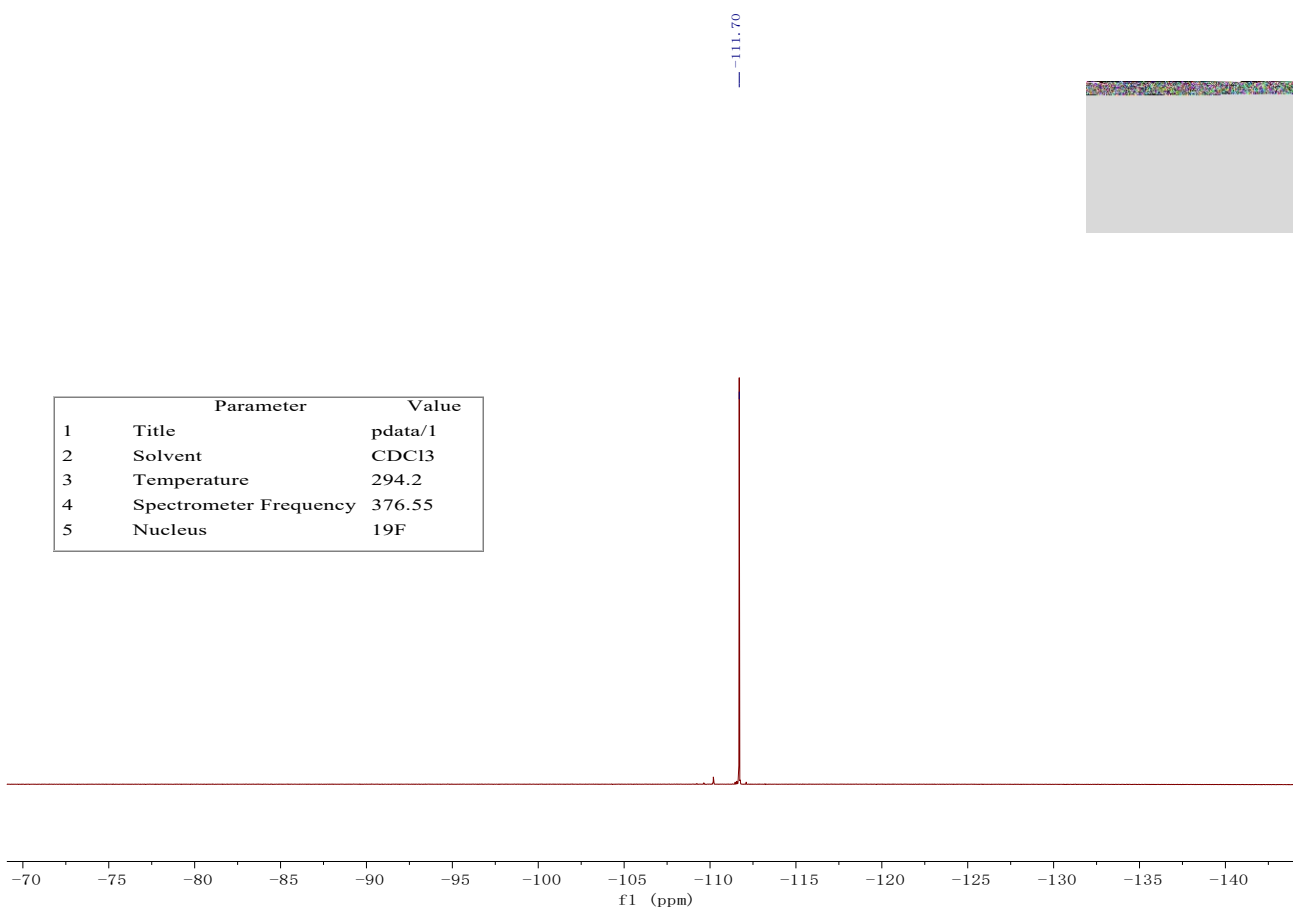


|   | Parameter              | Value          |
|---|------------------------|----------------|
| 1 | Title                  | pdata/1        |
| 2 | Solvent                | CDC13          |
| 3 | Temperature            | 294.0          |
| 4 | Spectrometer Frequency | 400.18         |
| 5 | Nucleus                | <sup>1</sup> H |

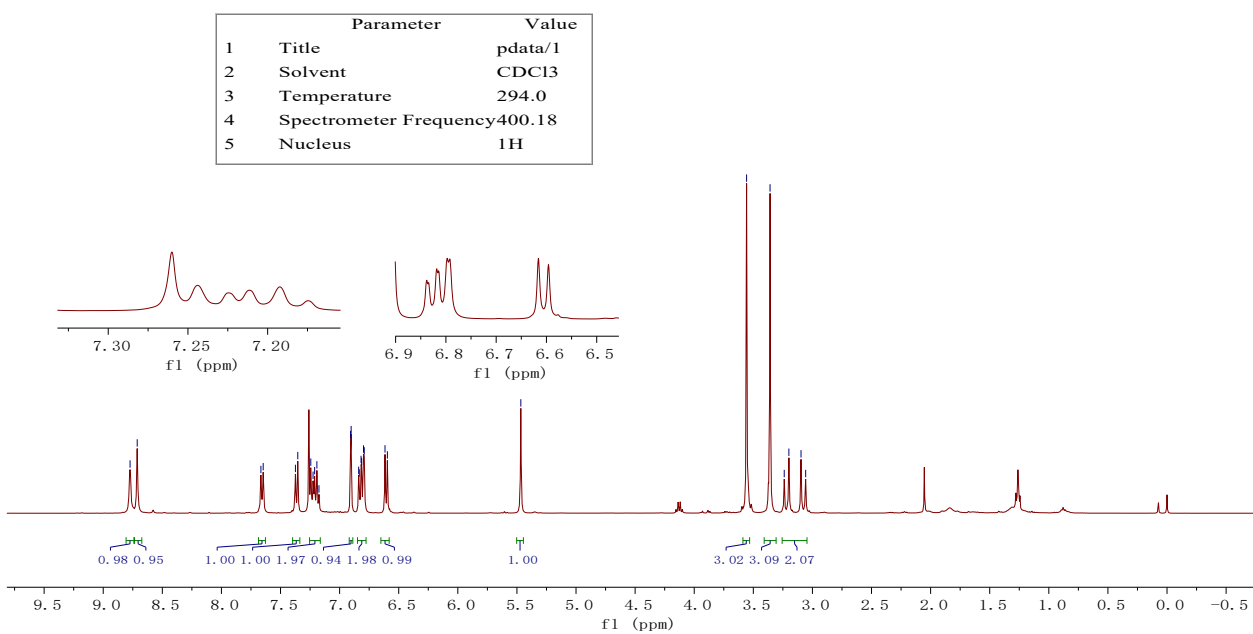
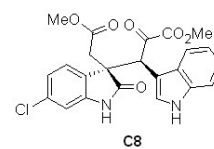


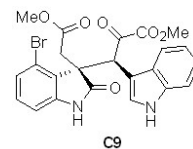
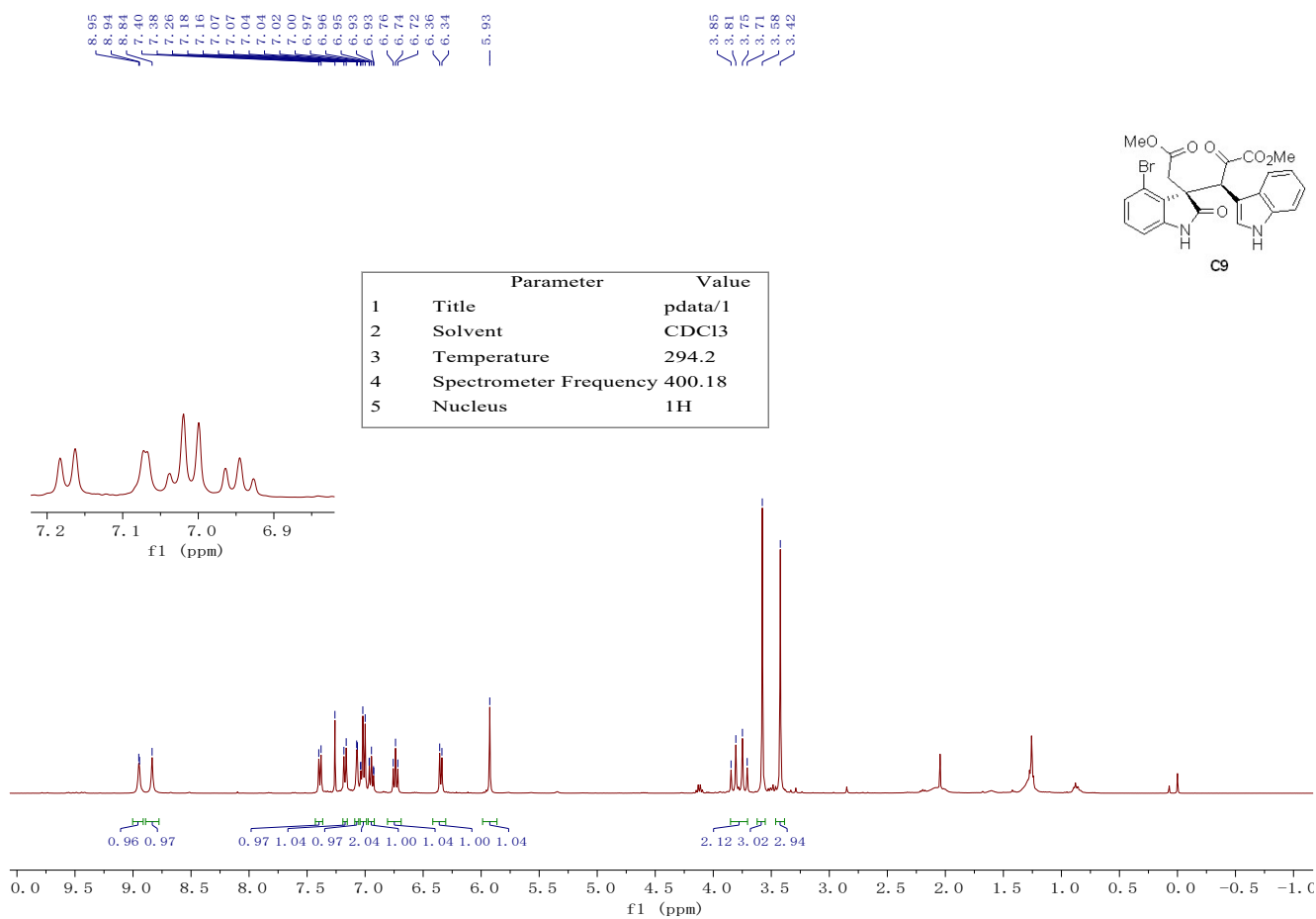
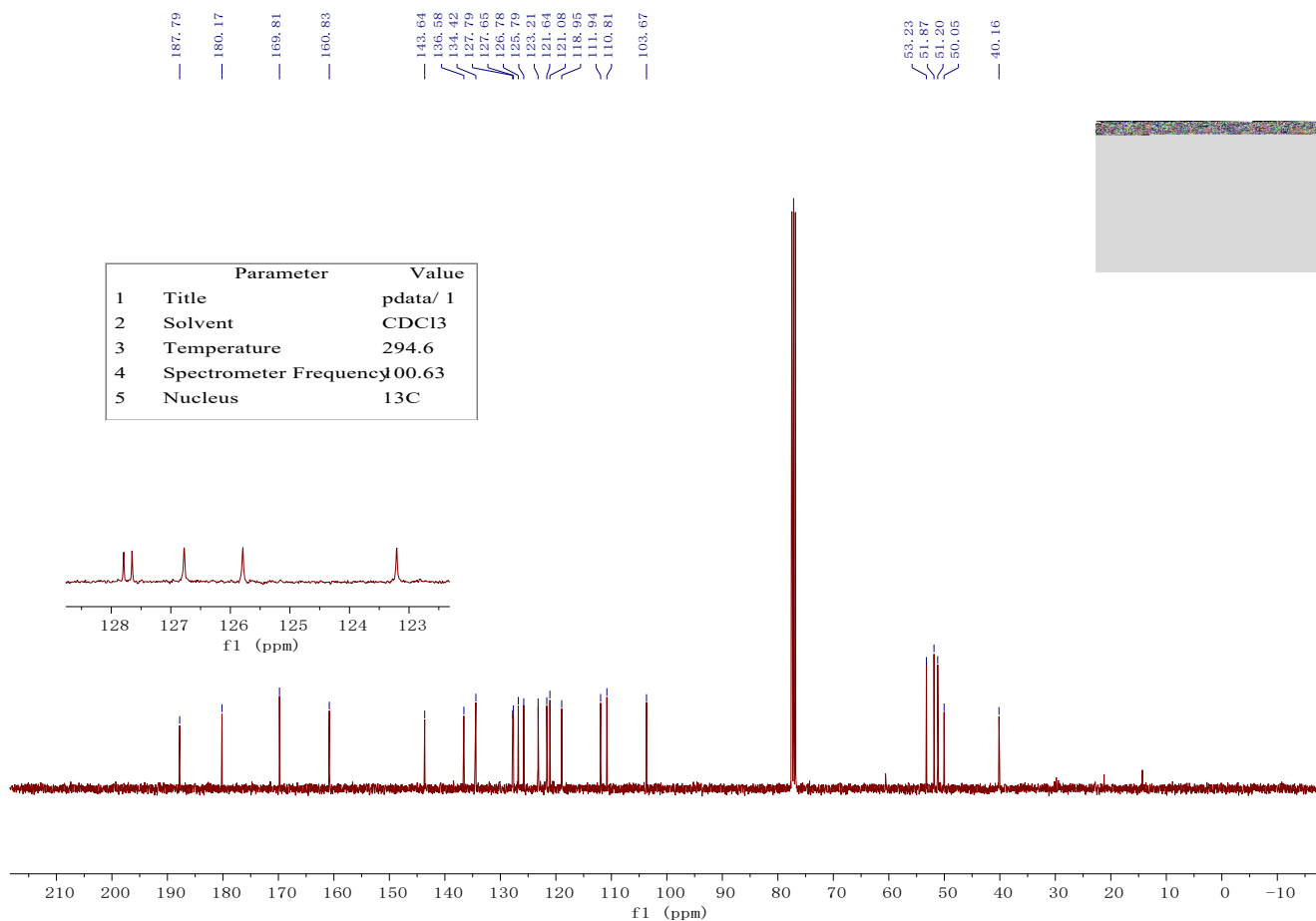
|   | Parameter              | Value           |
|---|------------------------|-----------------|
| 1 | Title                  | pdata/1         |
| 2 | Solvent                | CDC13           |
| 3 | Temperature            | 294.5           |
| 4 | Spectrometer Frequency | 100.63          |
| 5 | Nucleus                | <sup>13</sup> C |

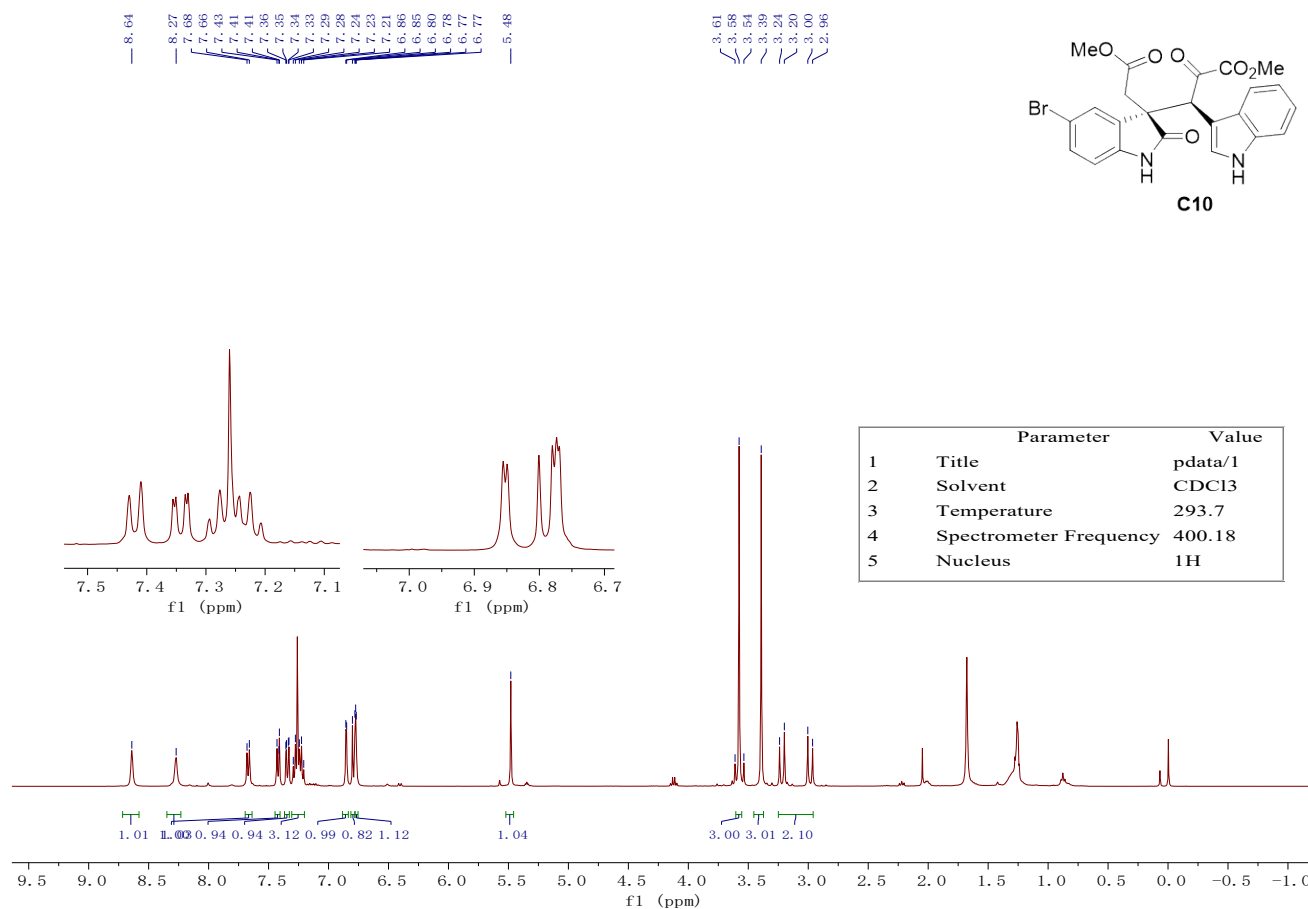
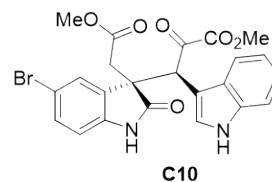
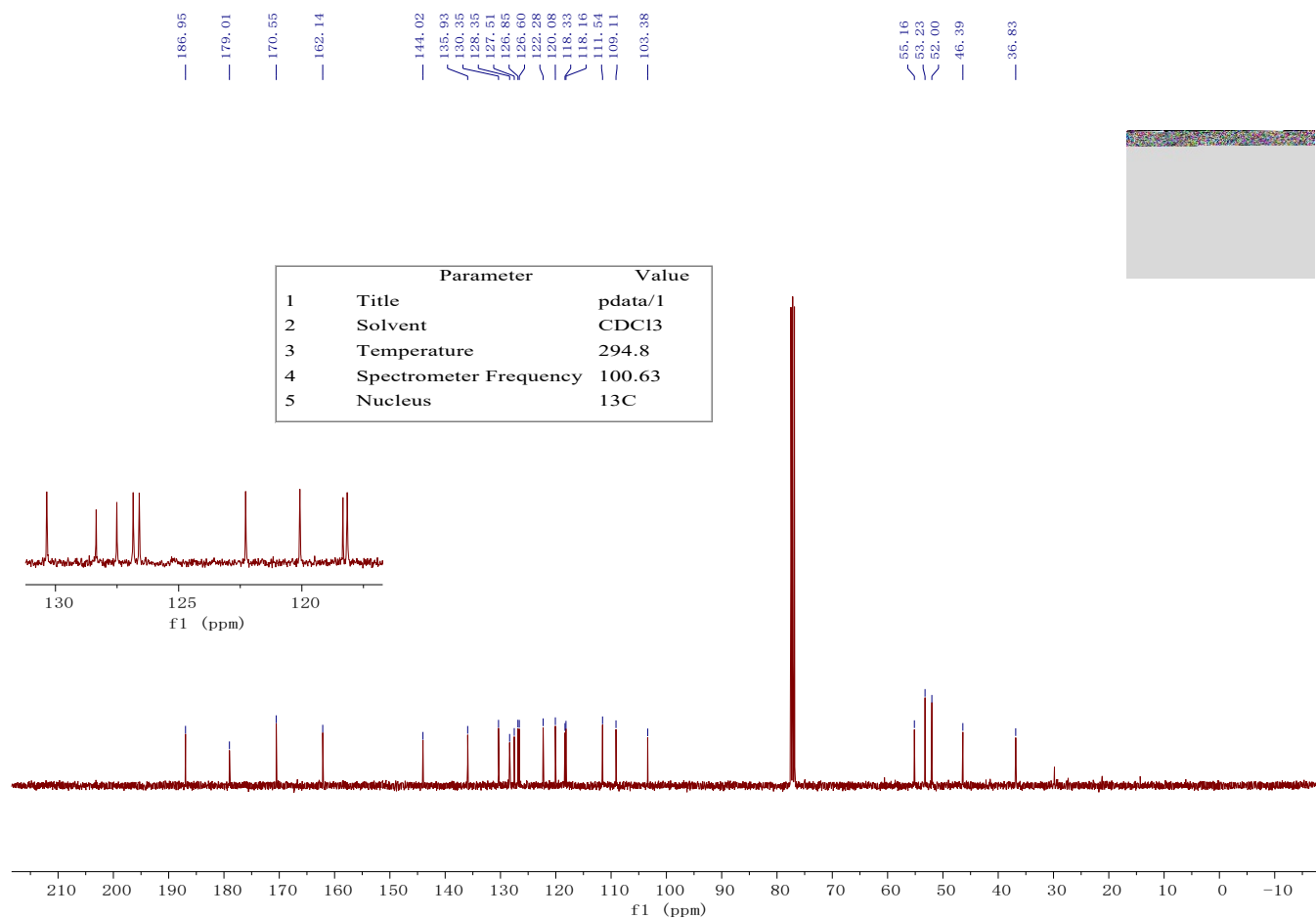


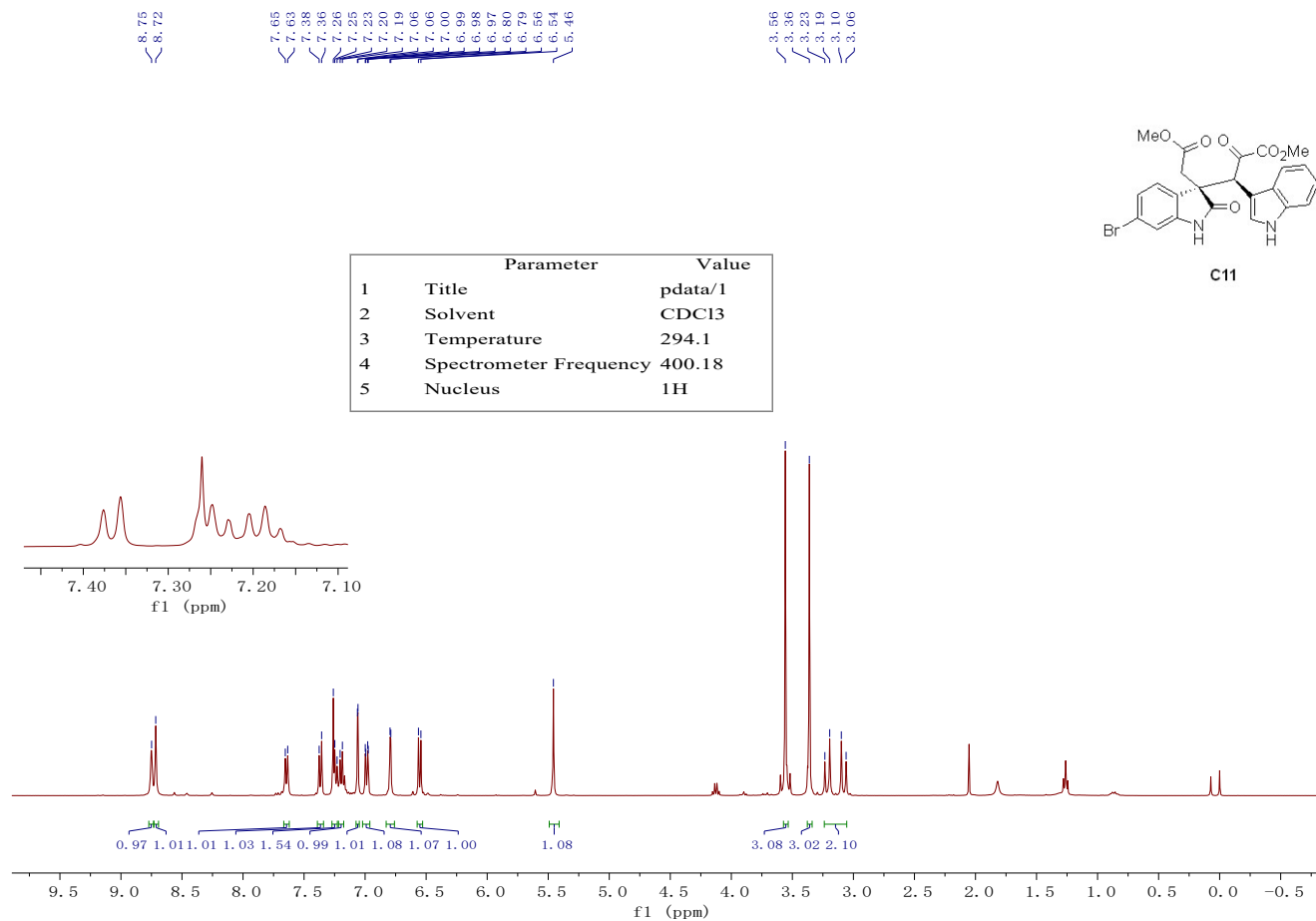
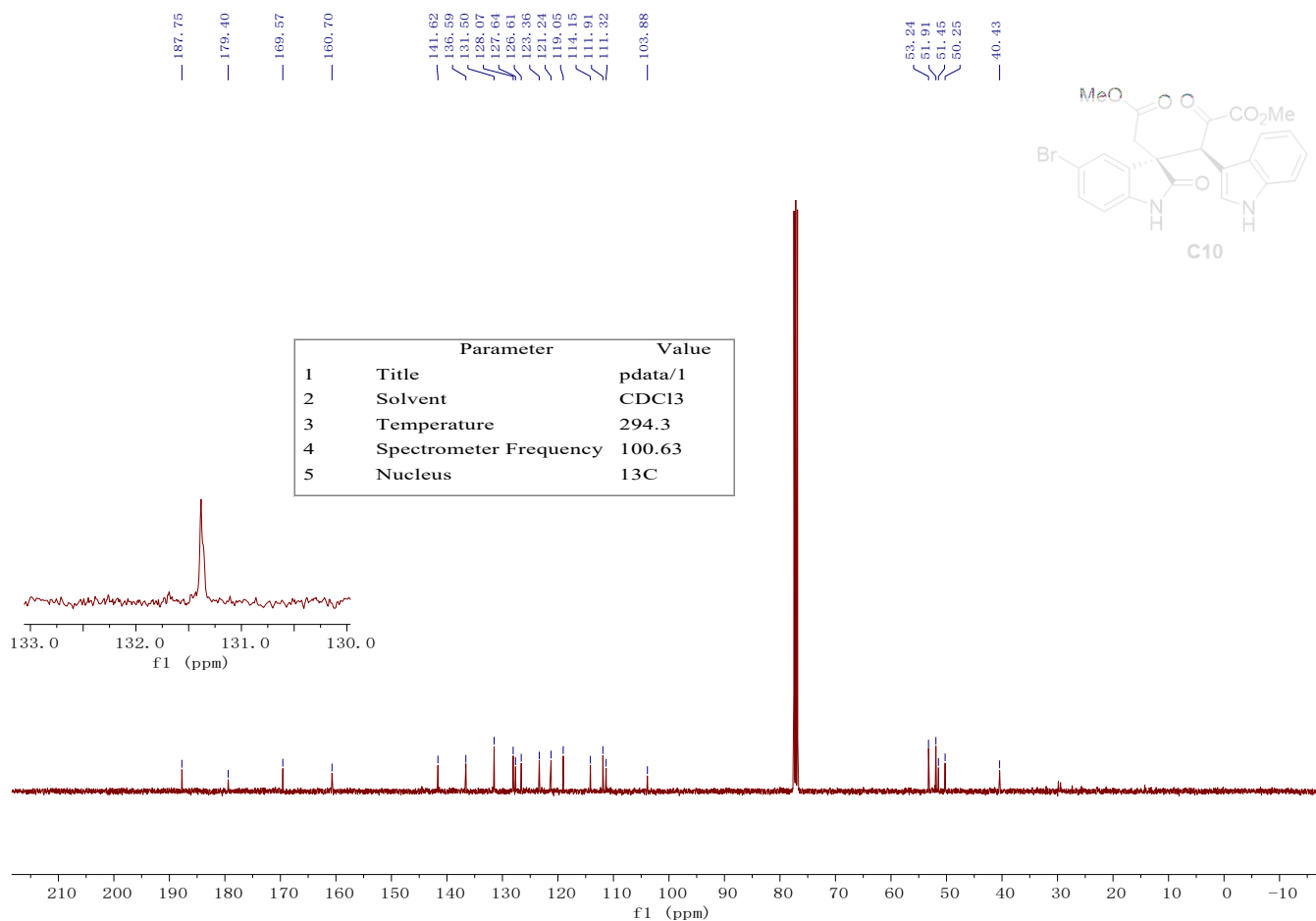


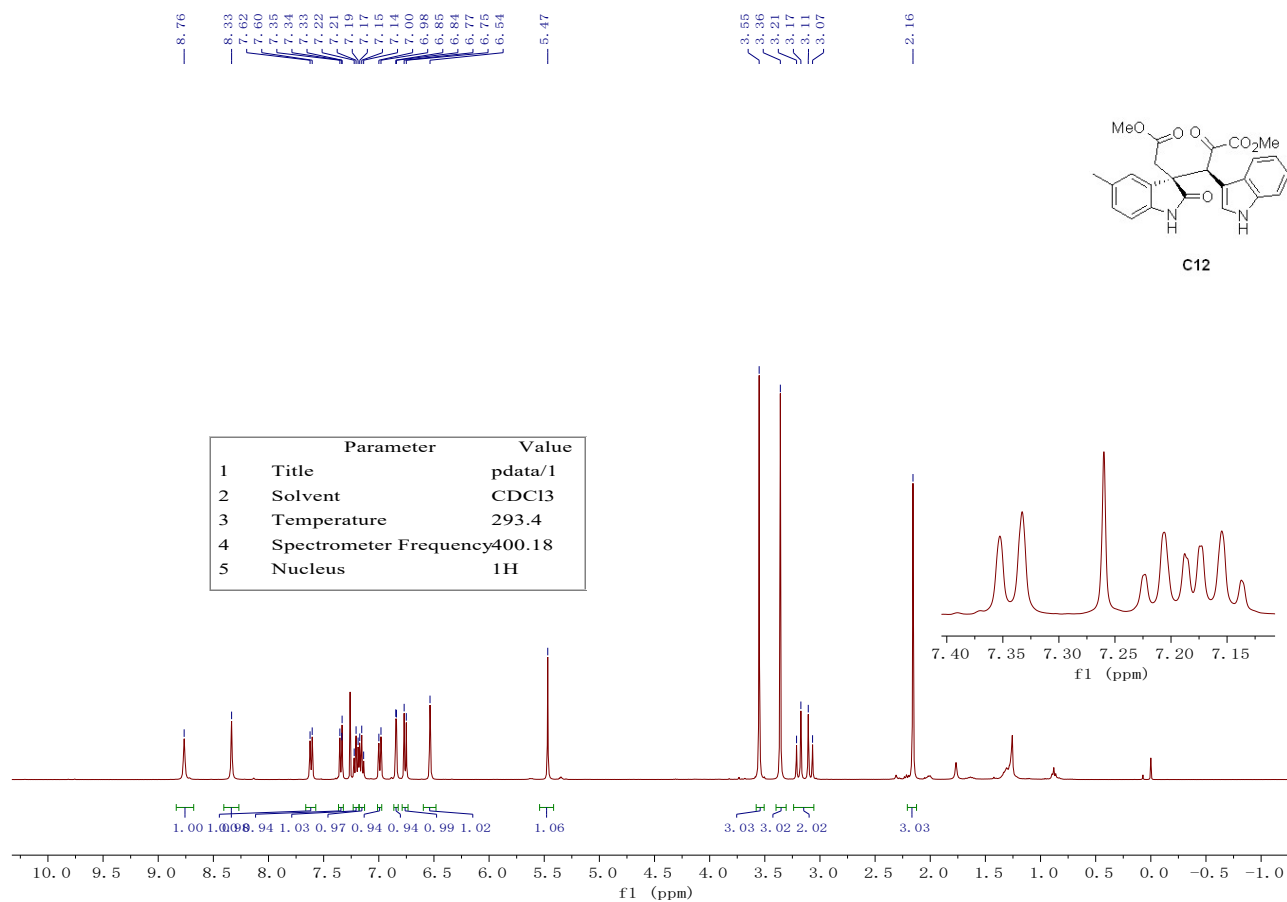
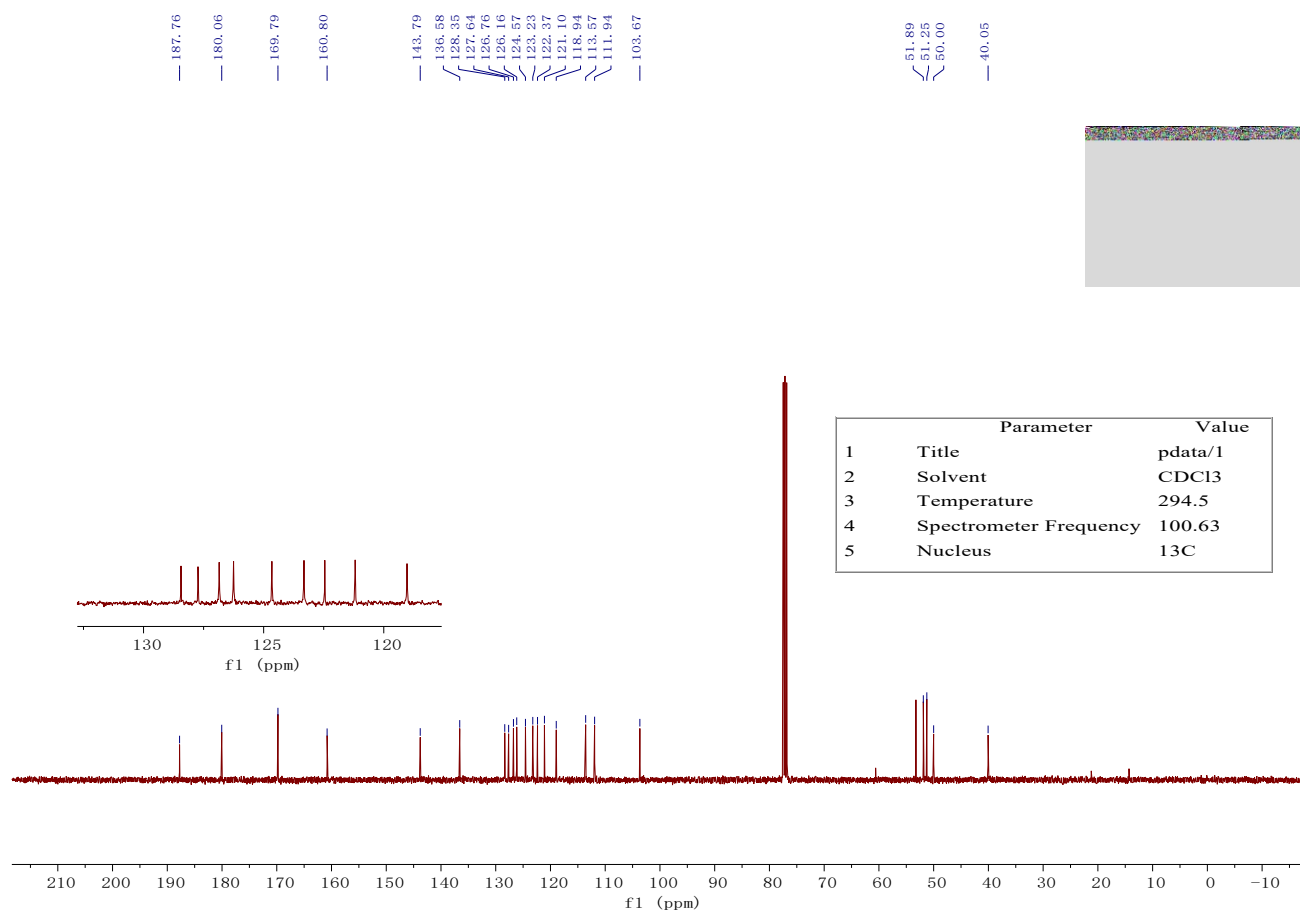
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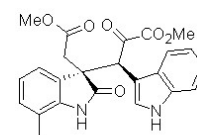
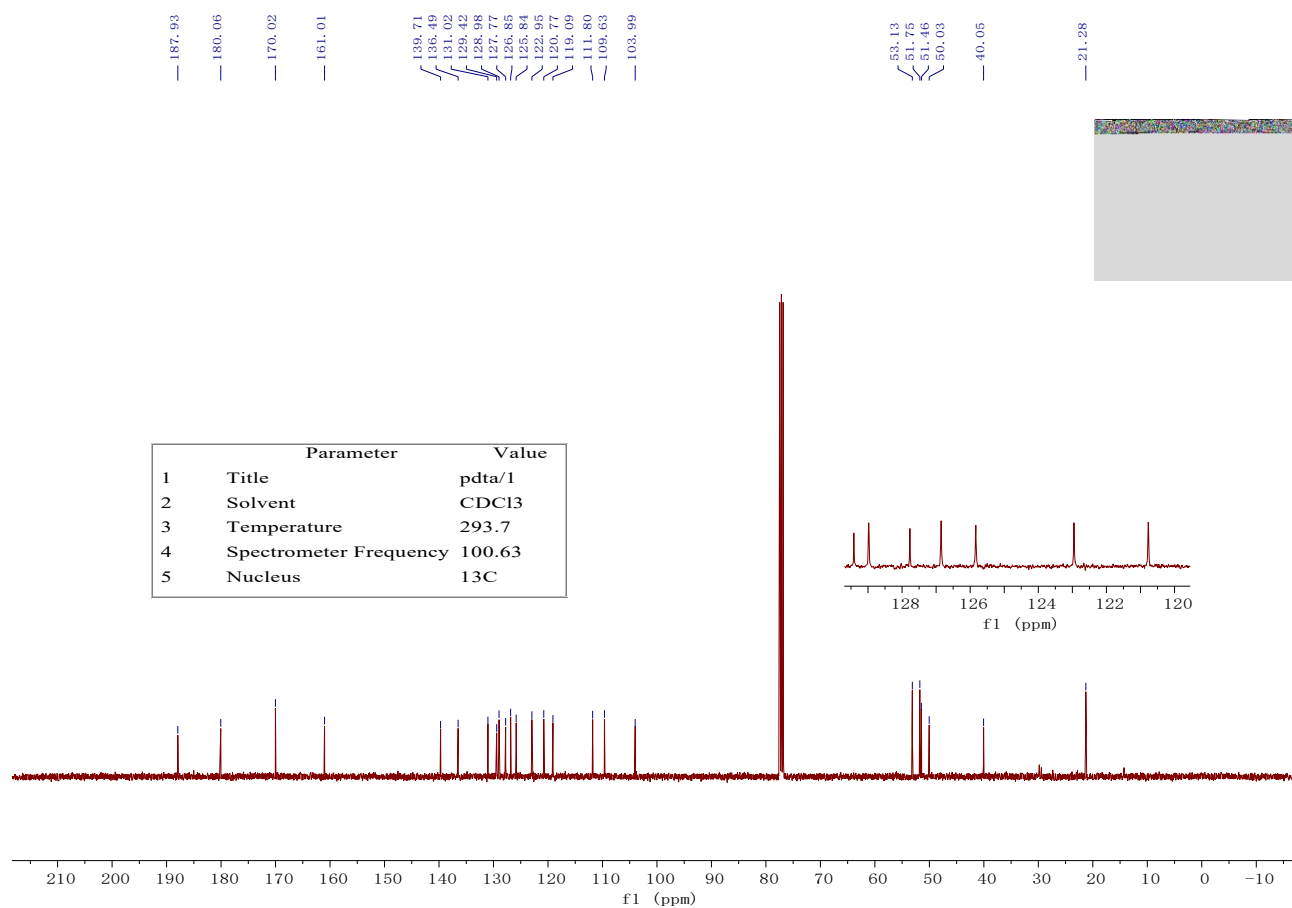




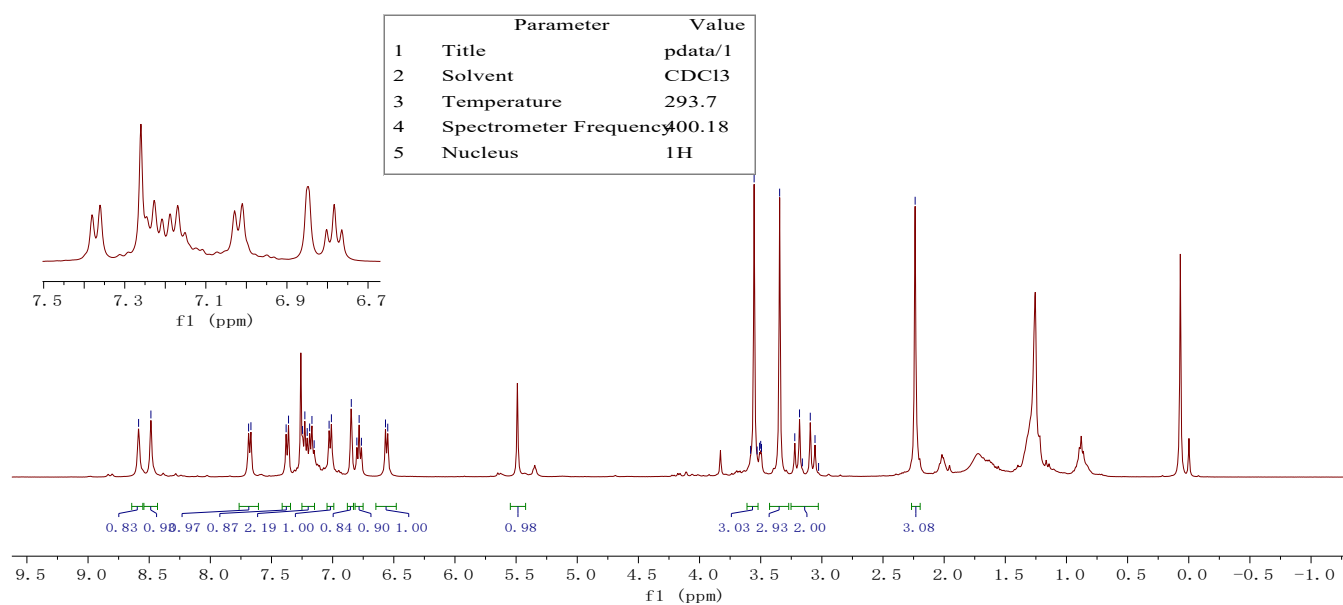


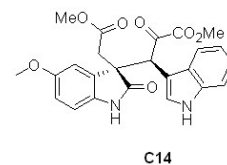
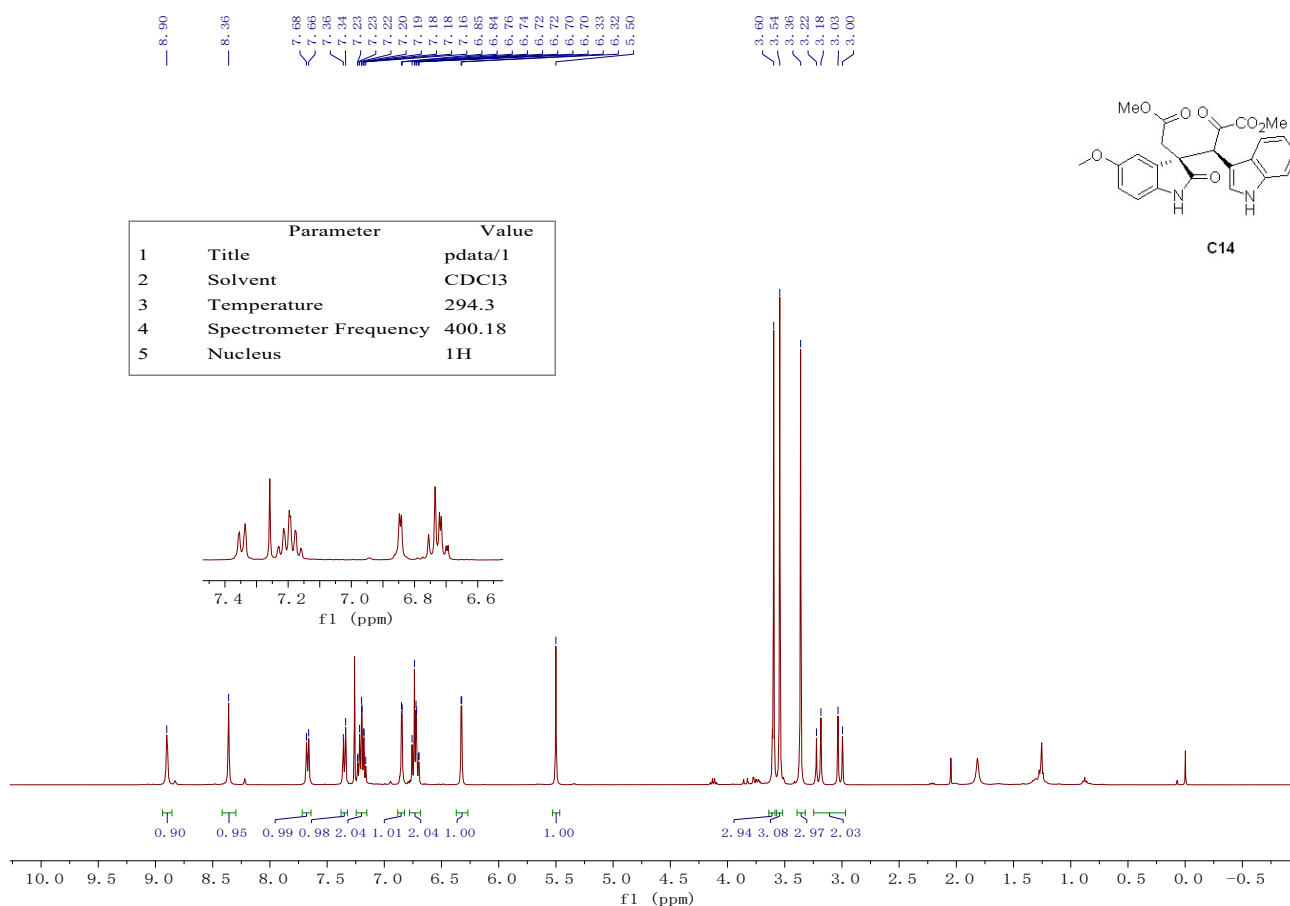
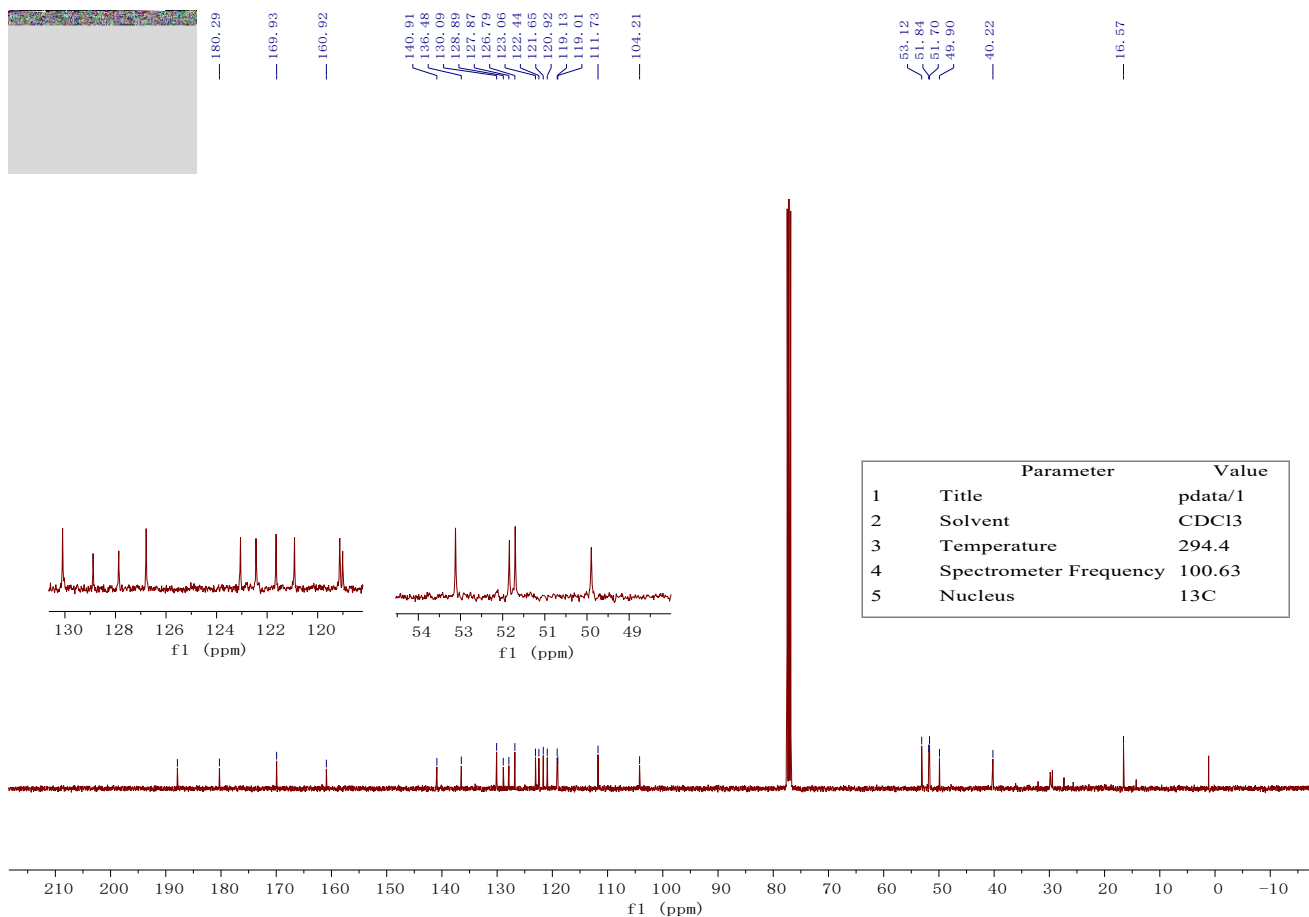


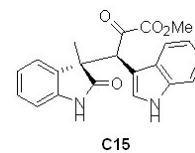
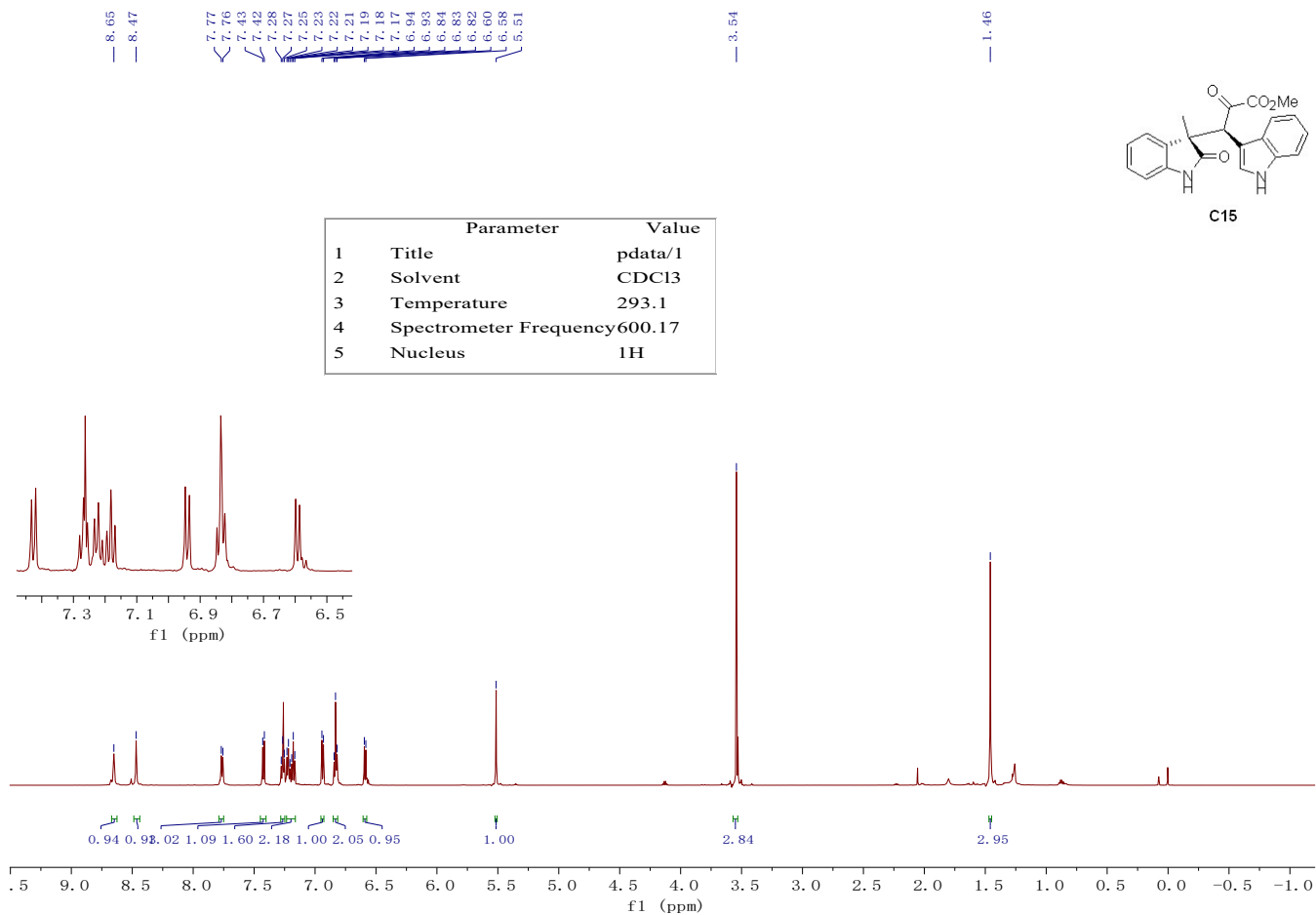
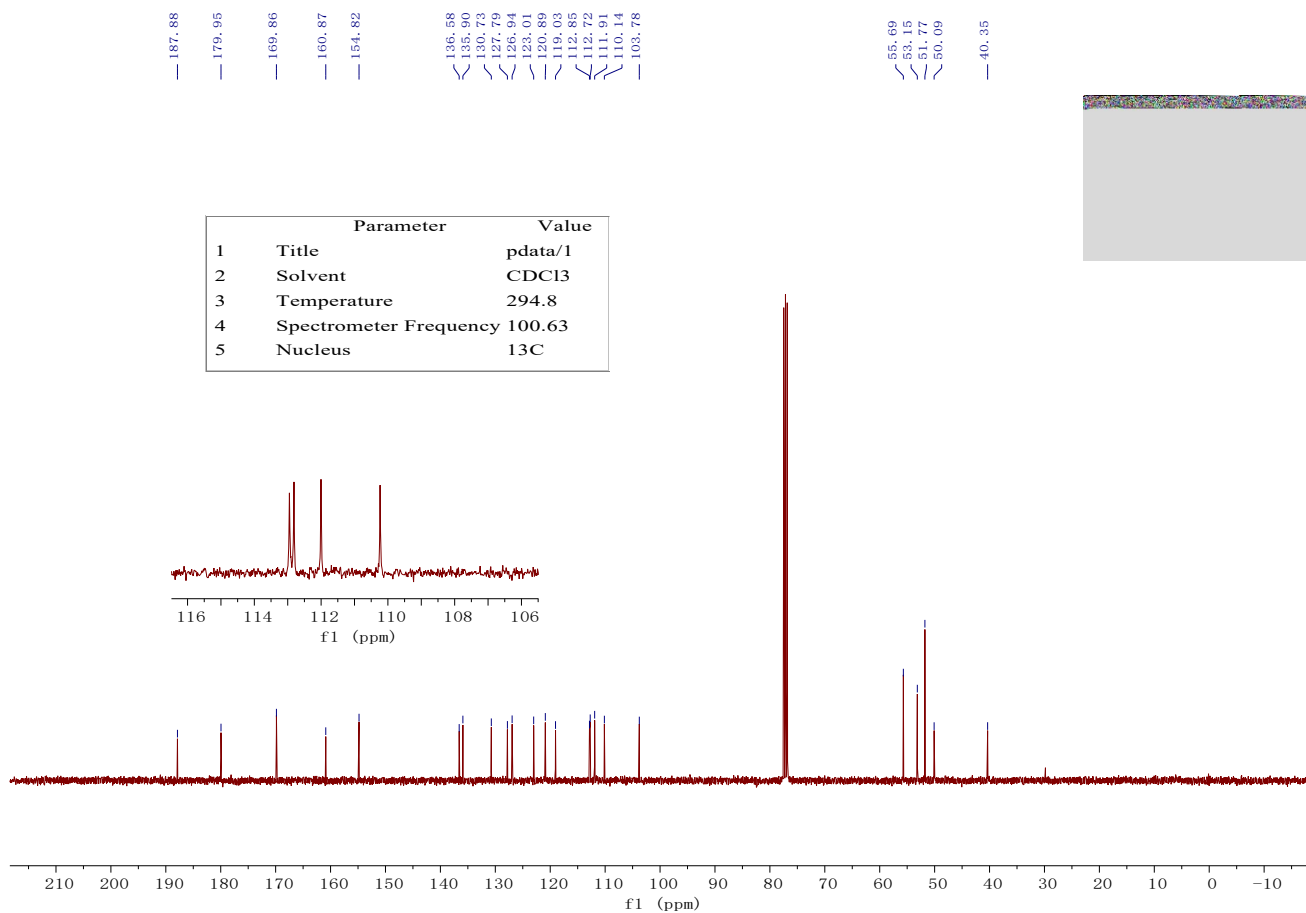


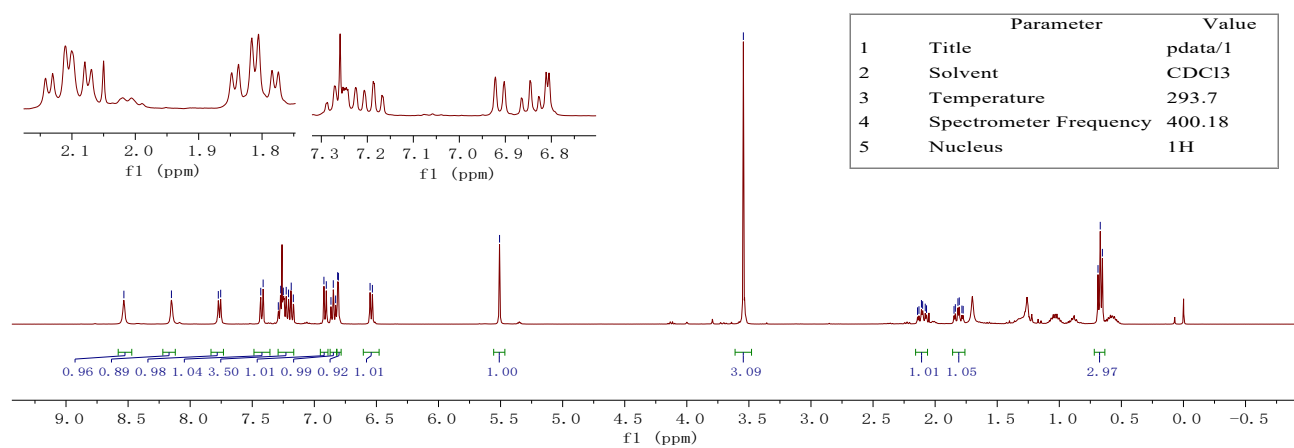
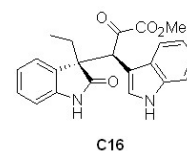
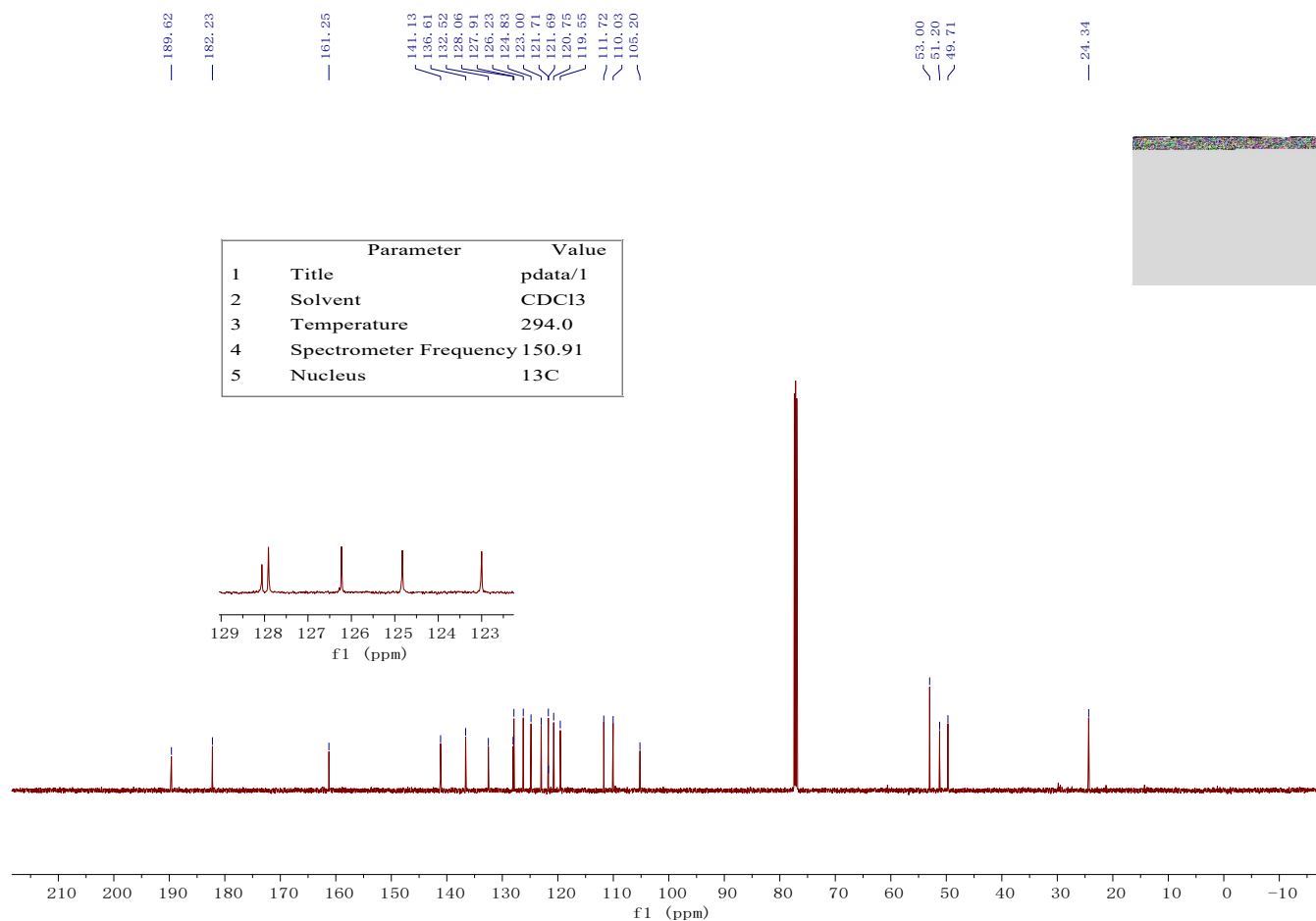


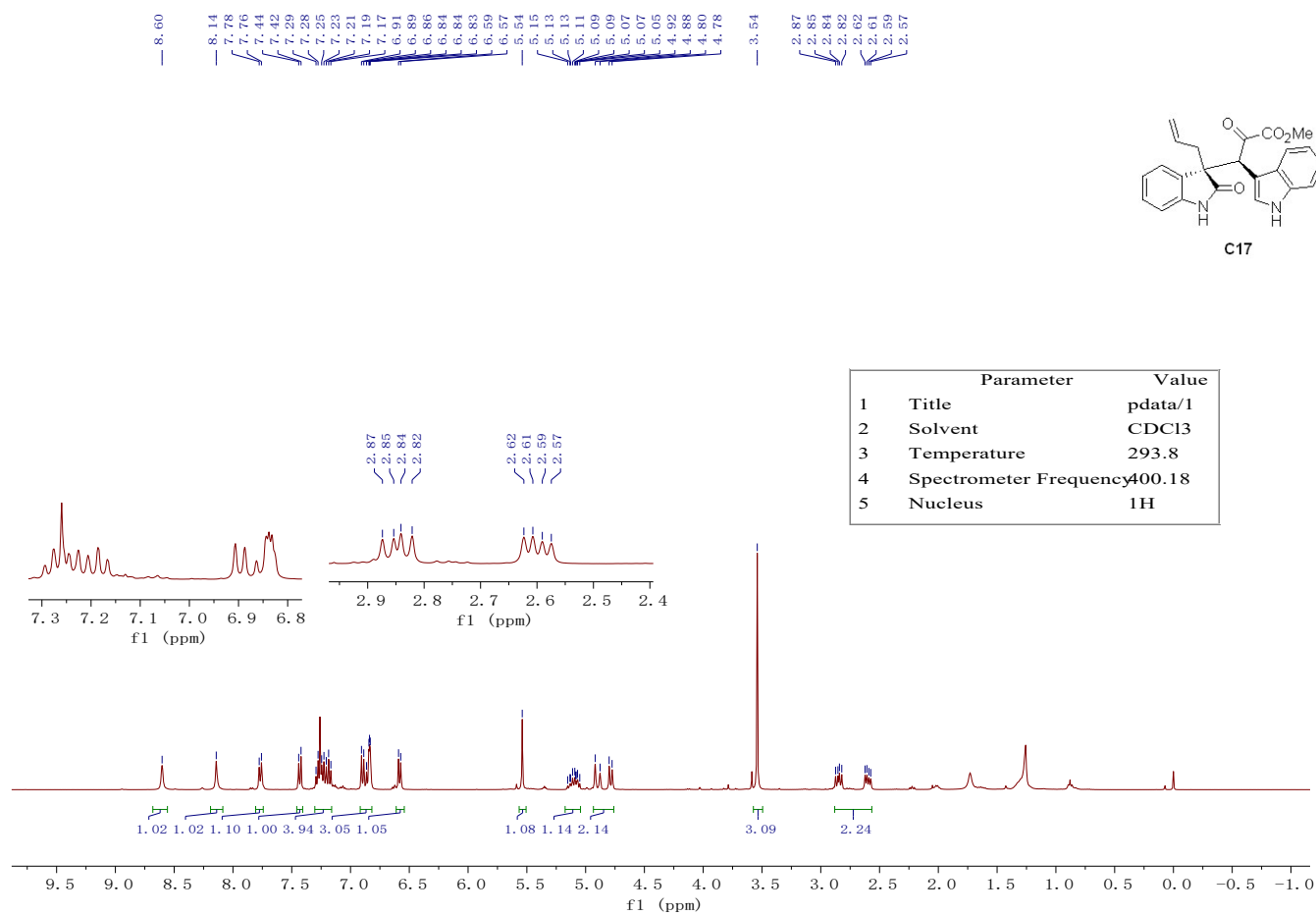
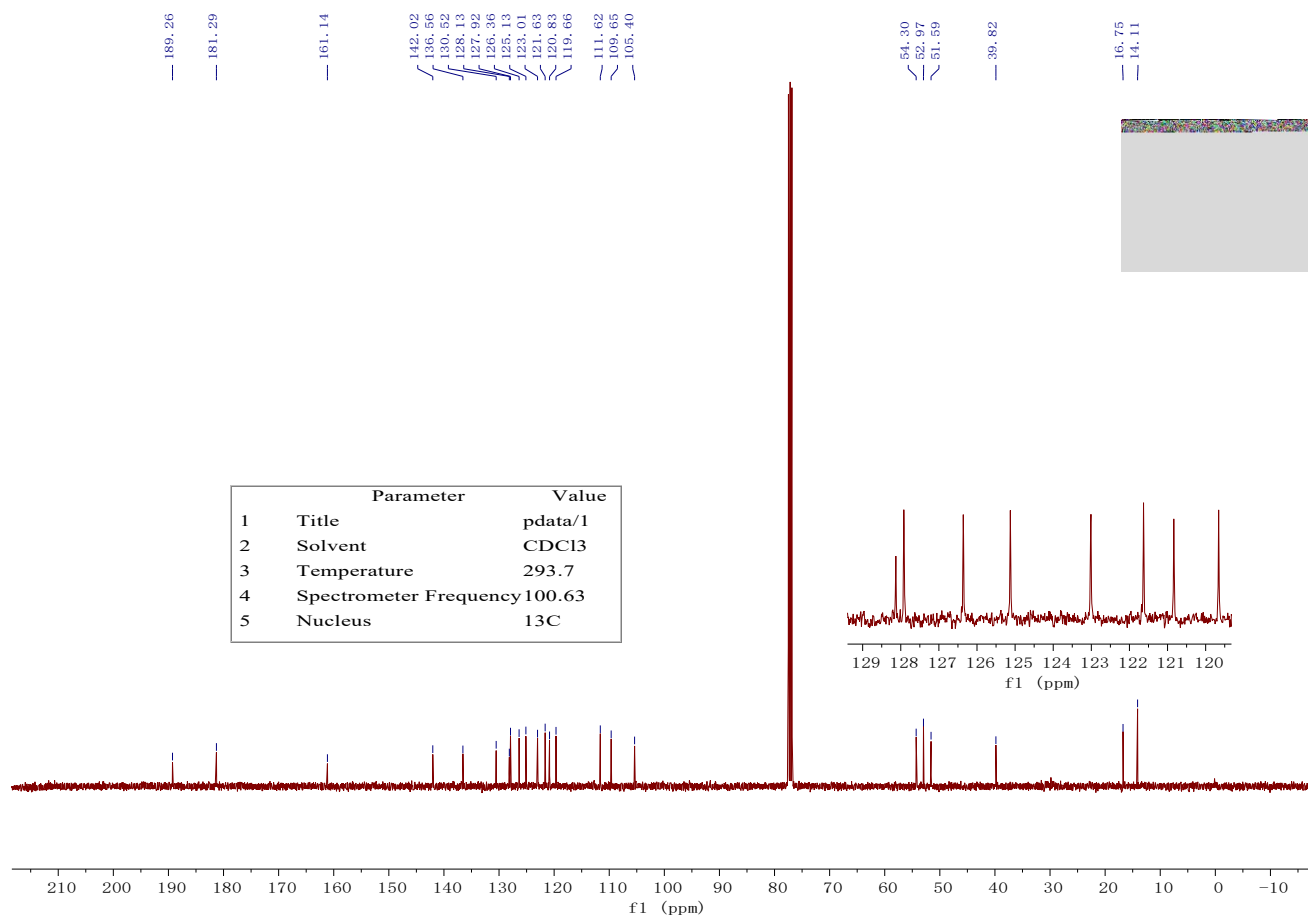
**C13**

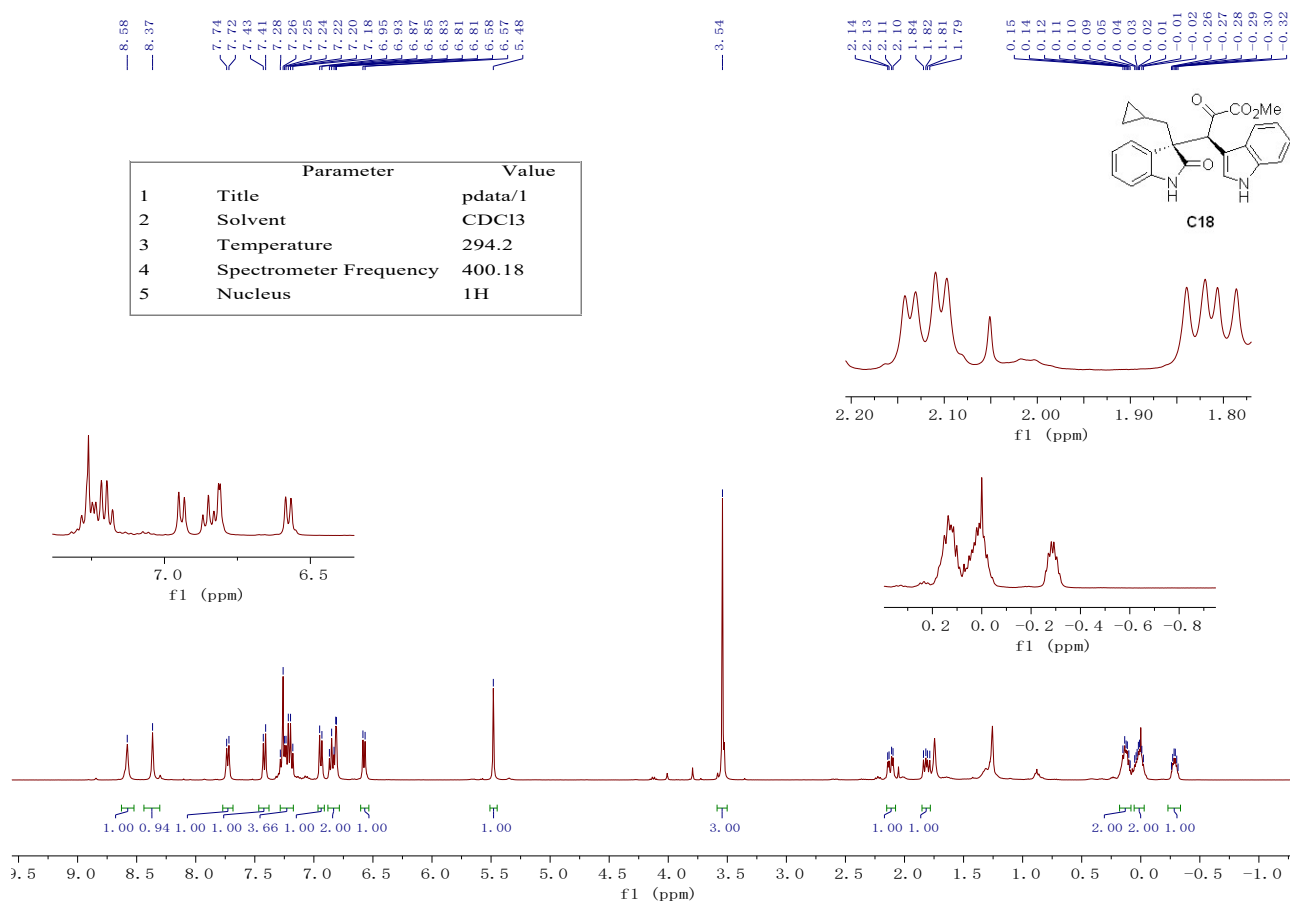
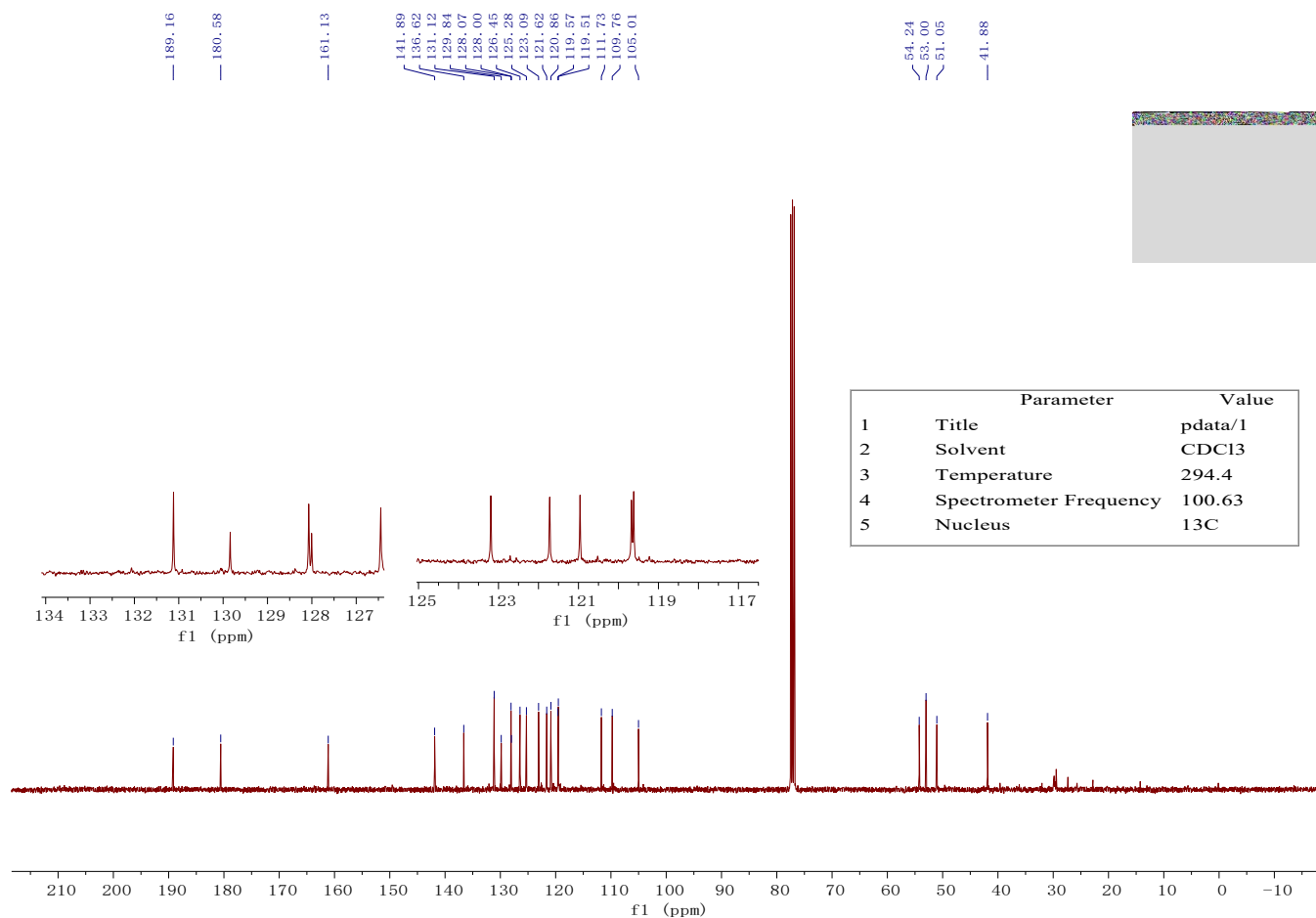


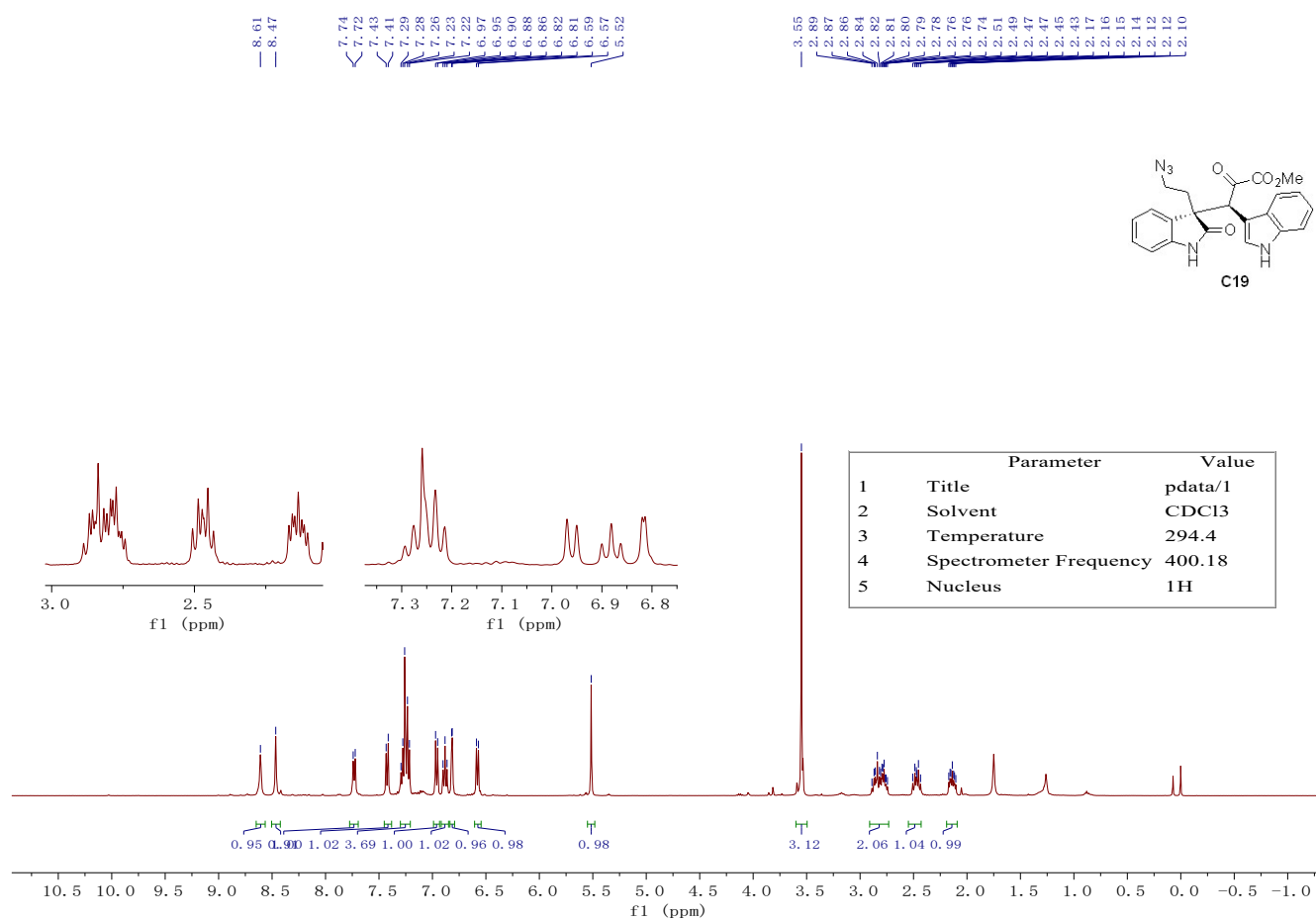
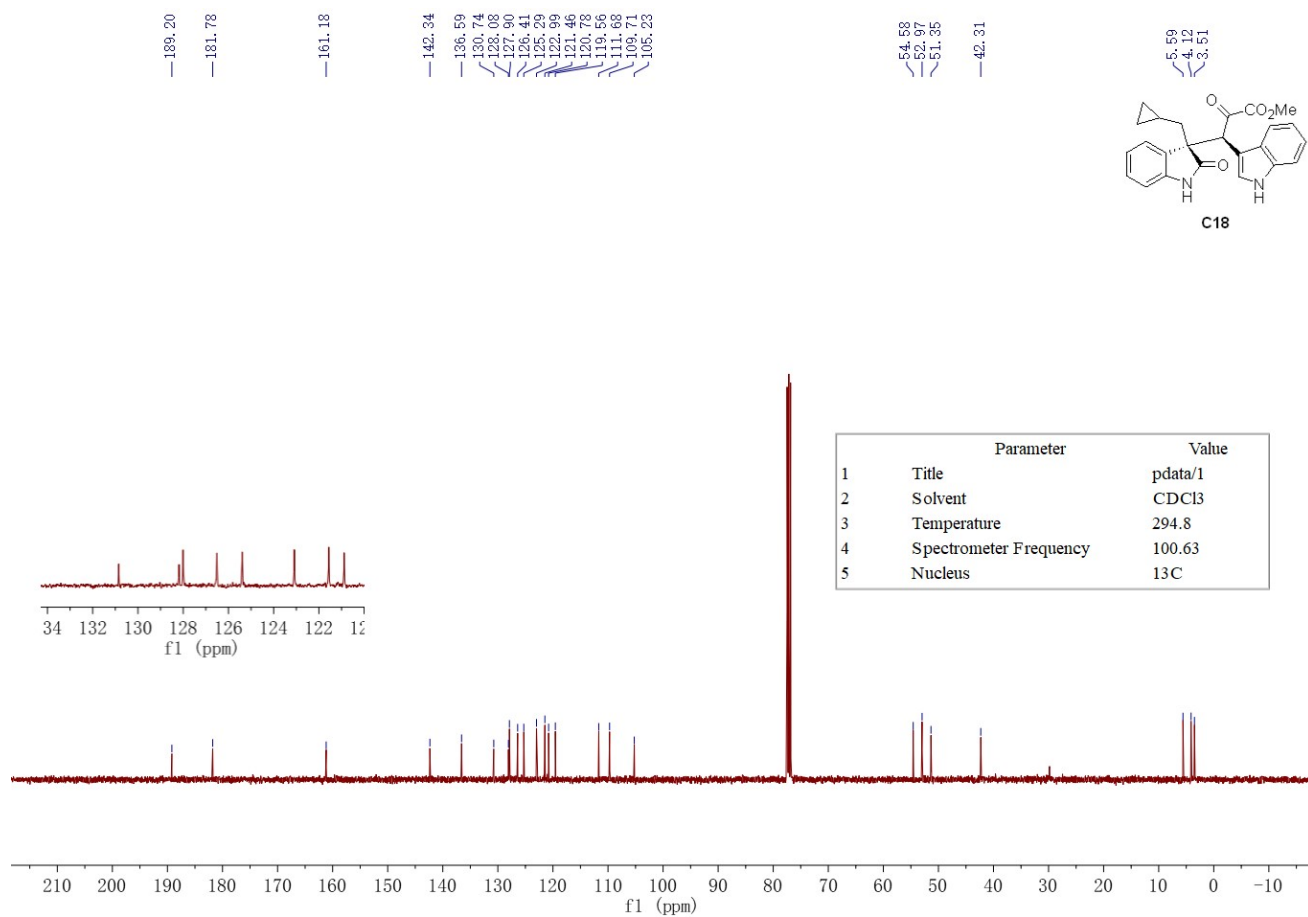


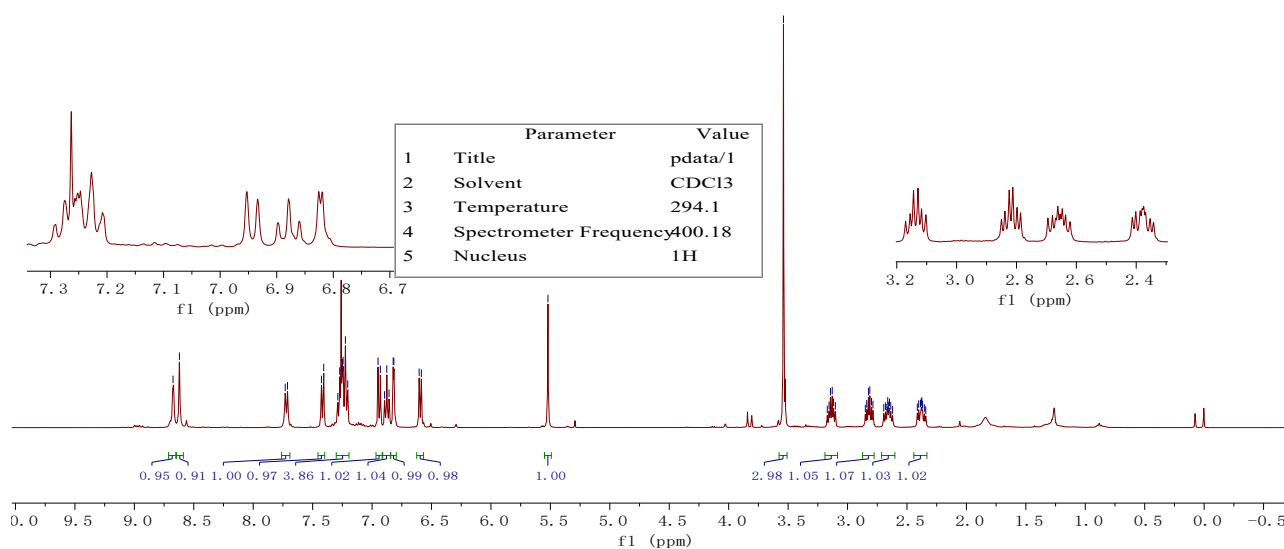
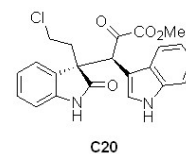
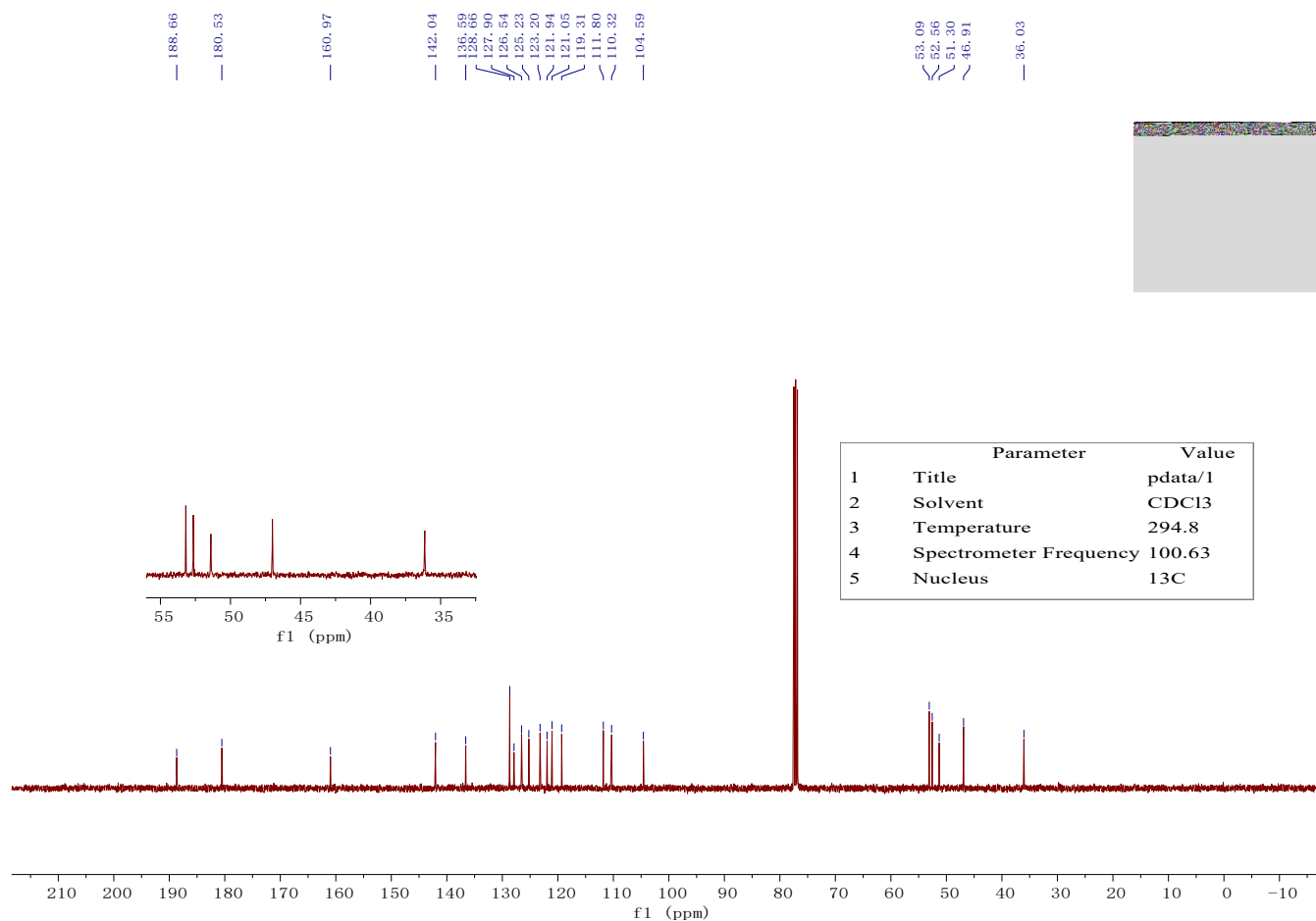


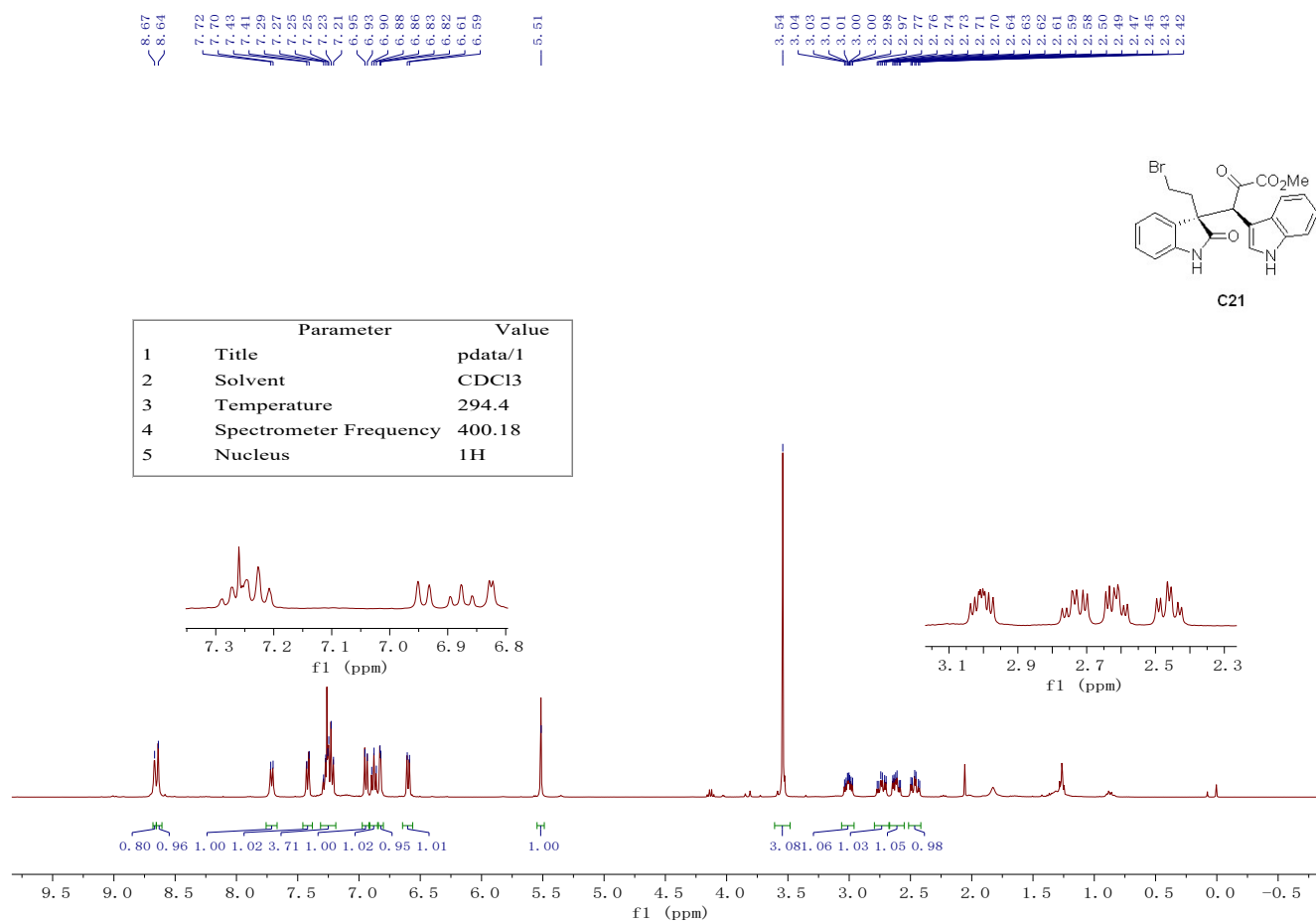
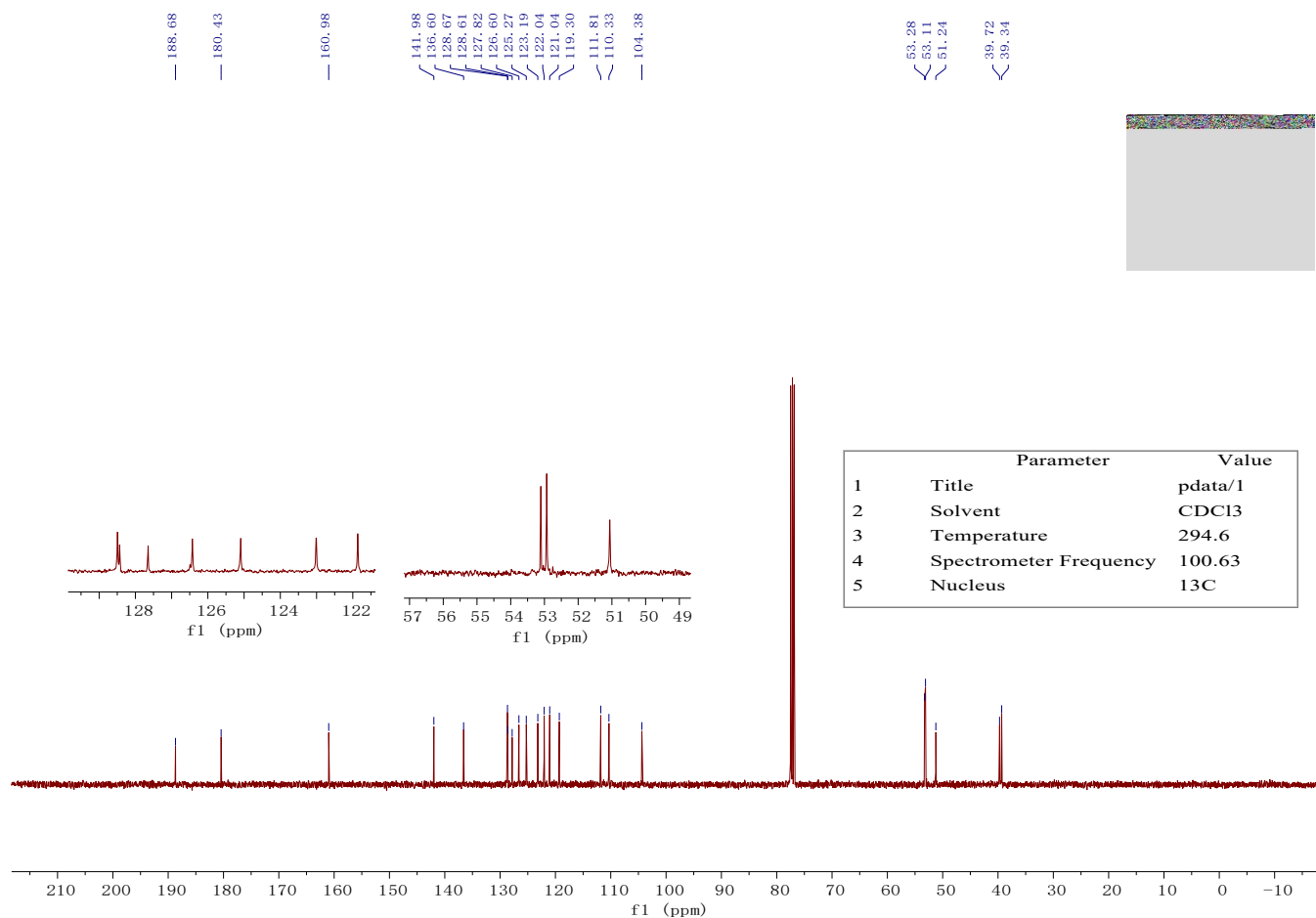


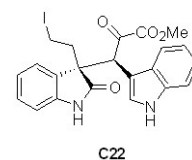
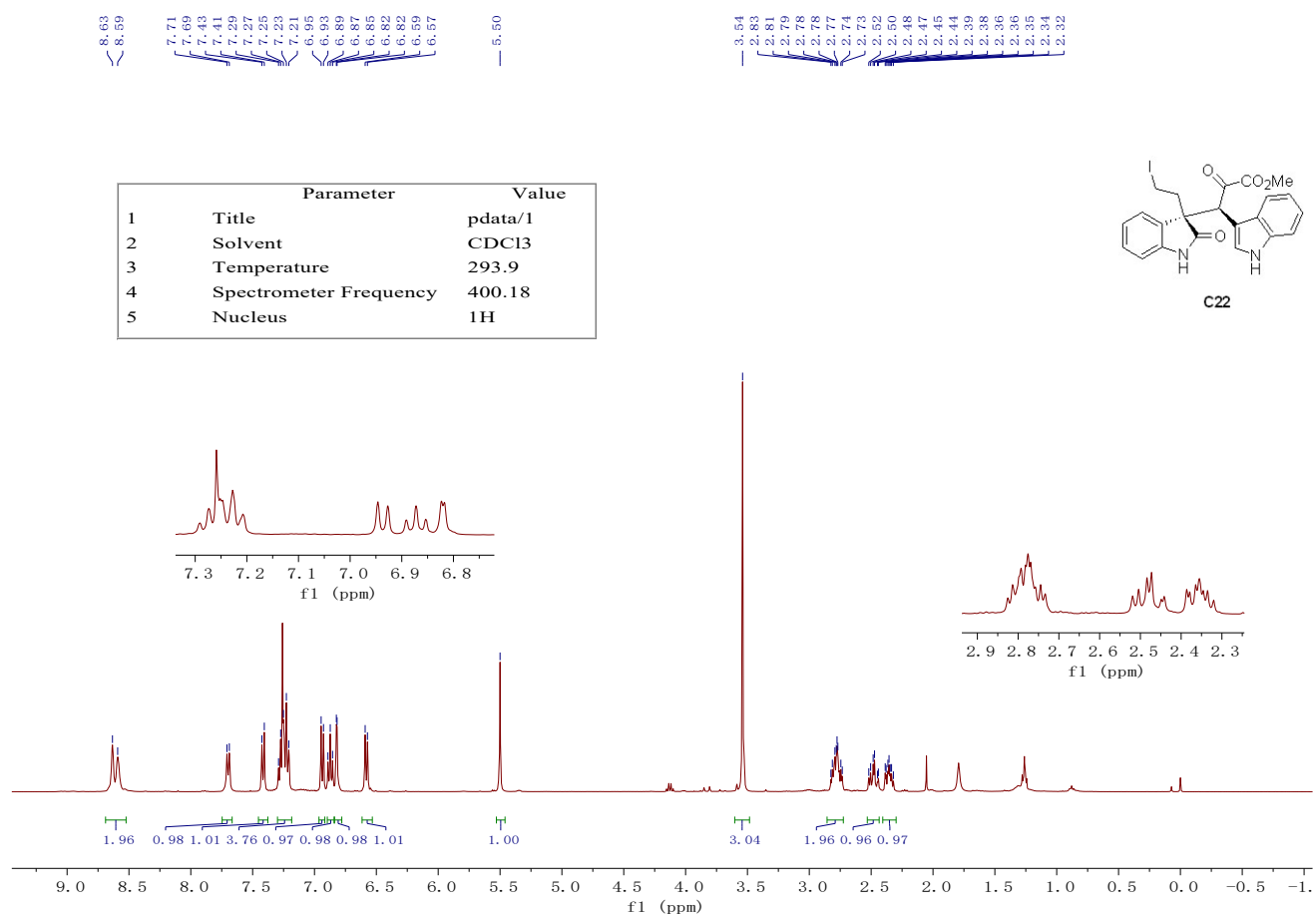
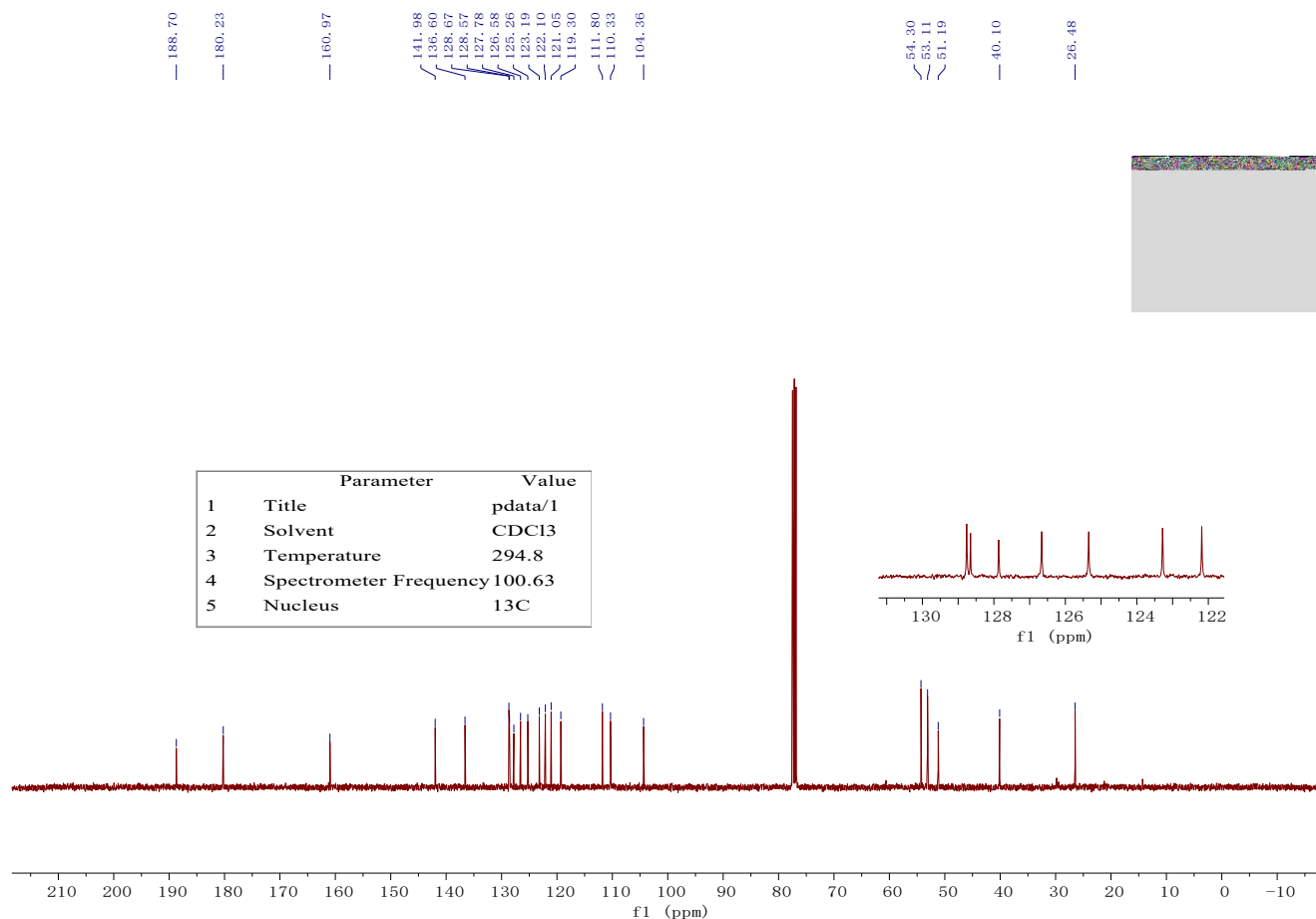


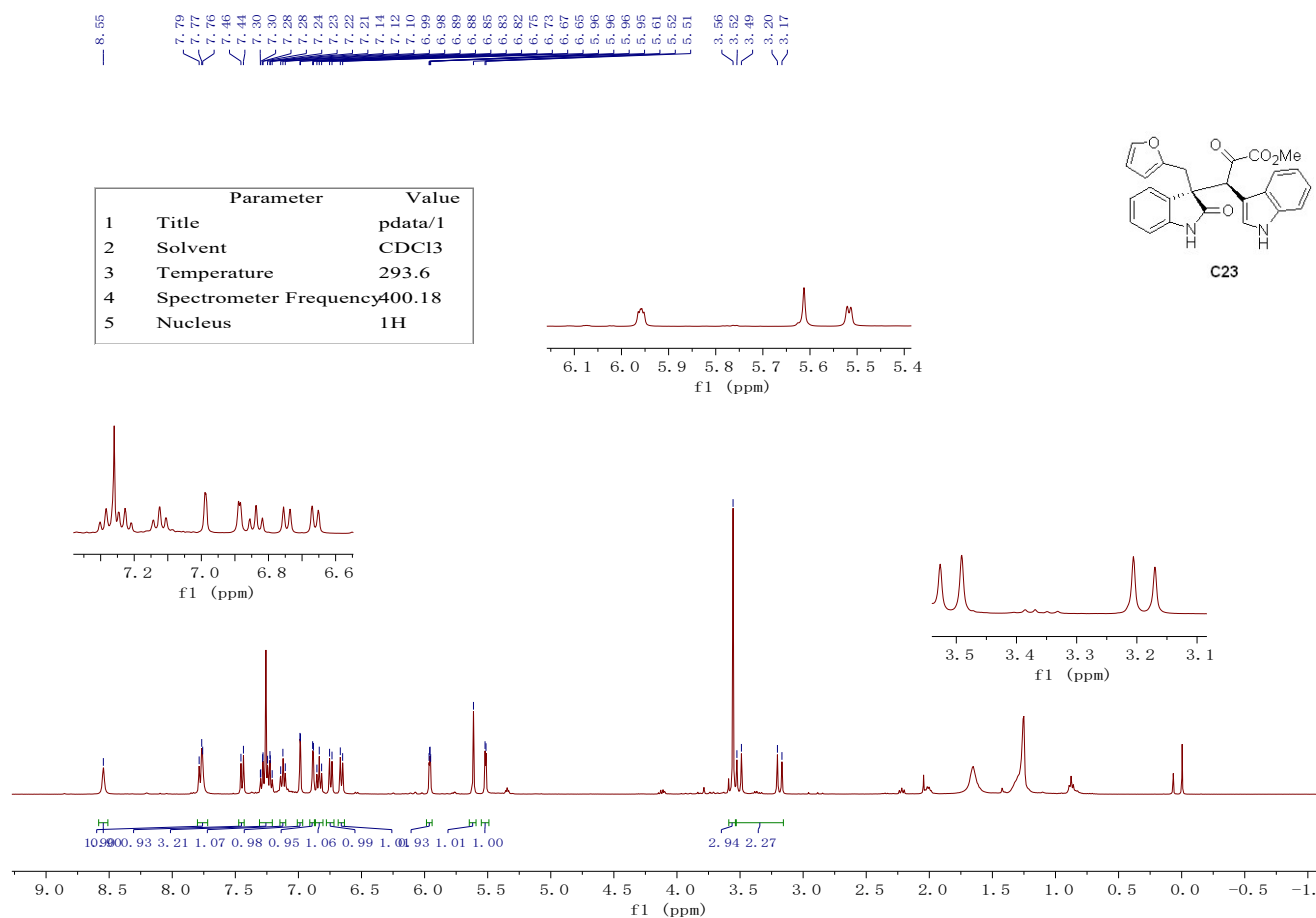
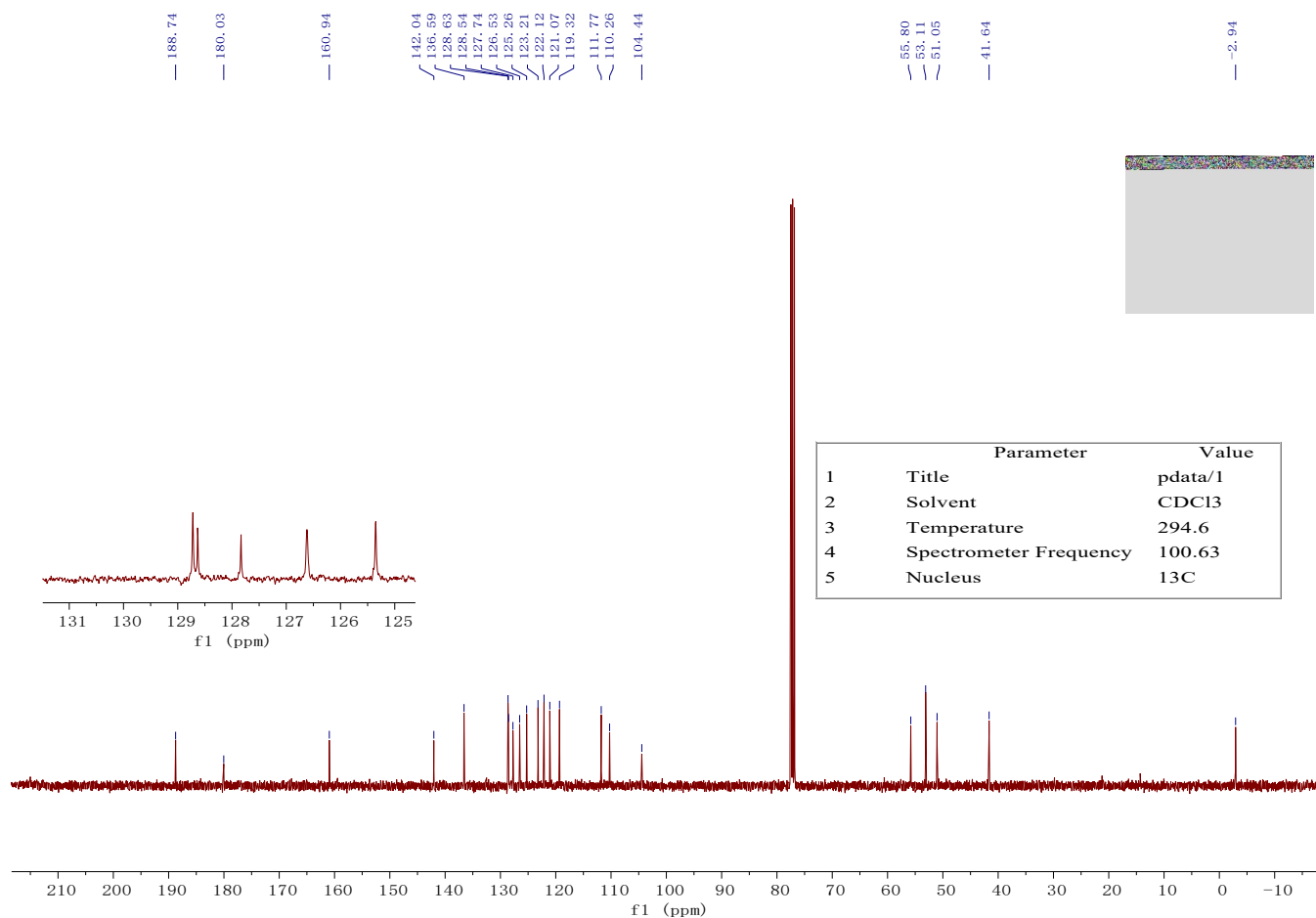


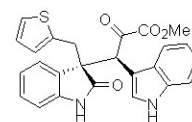
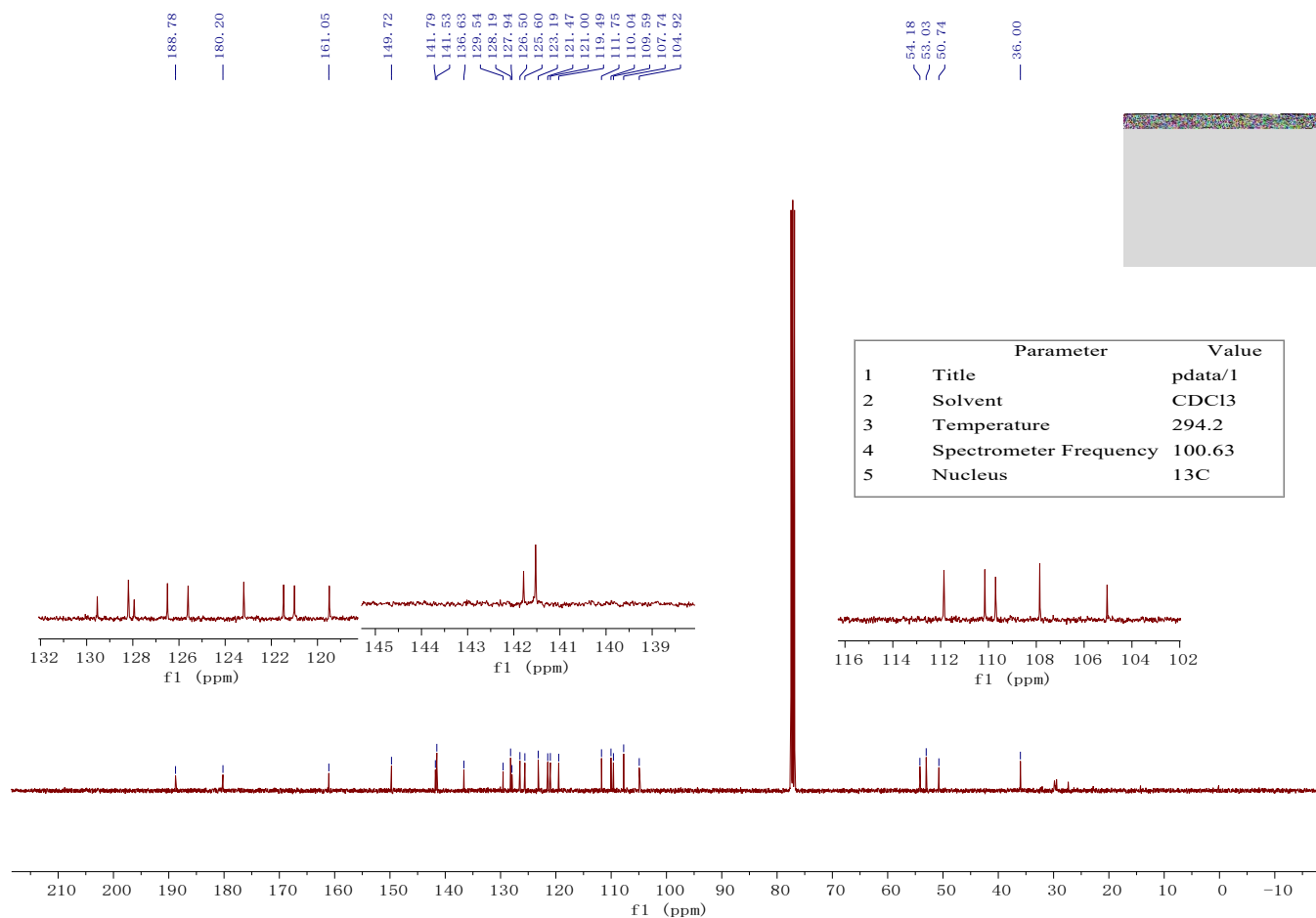




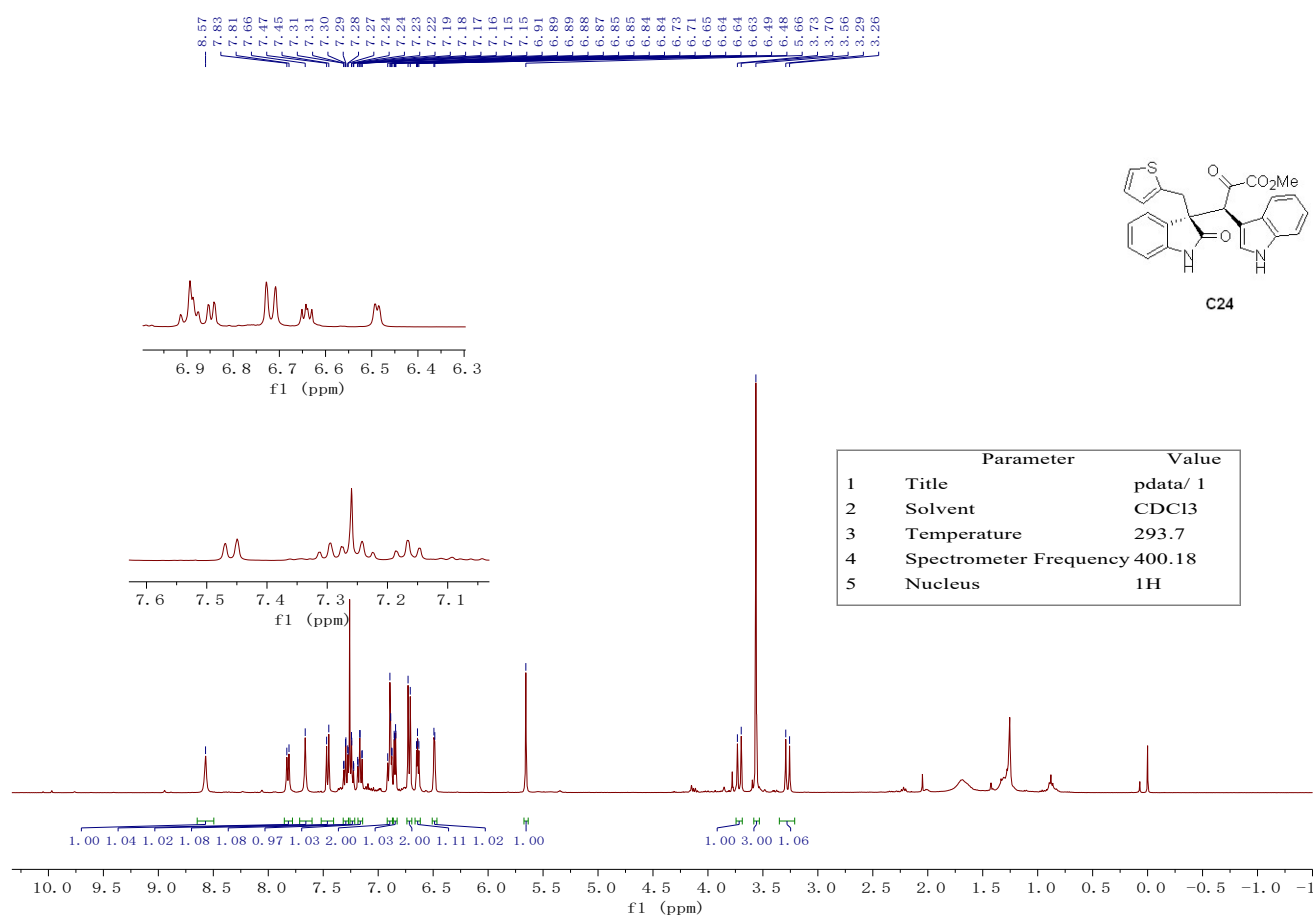


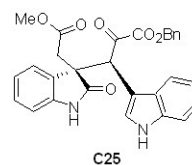
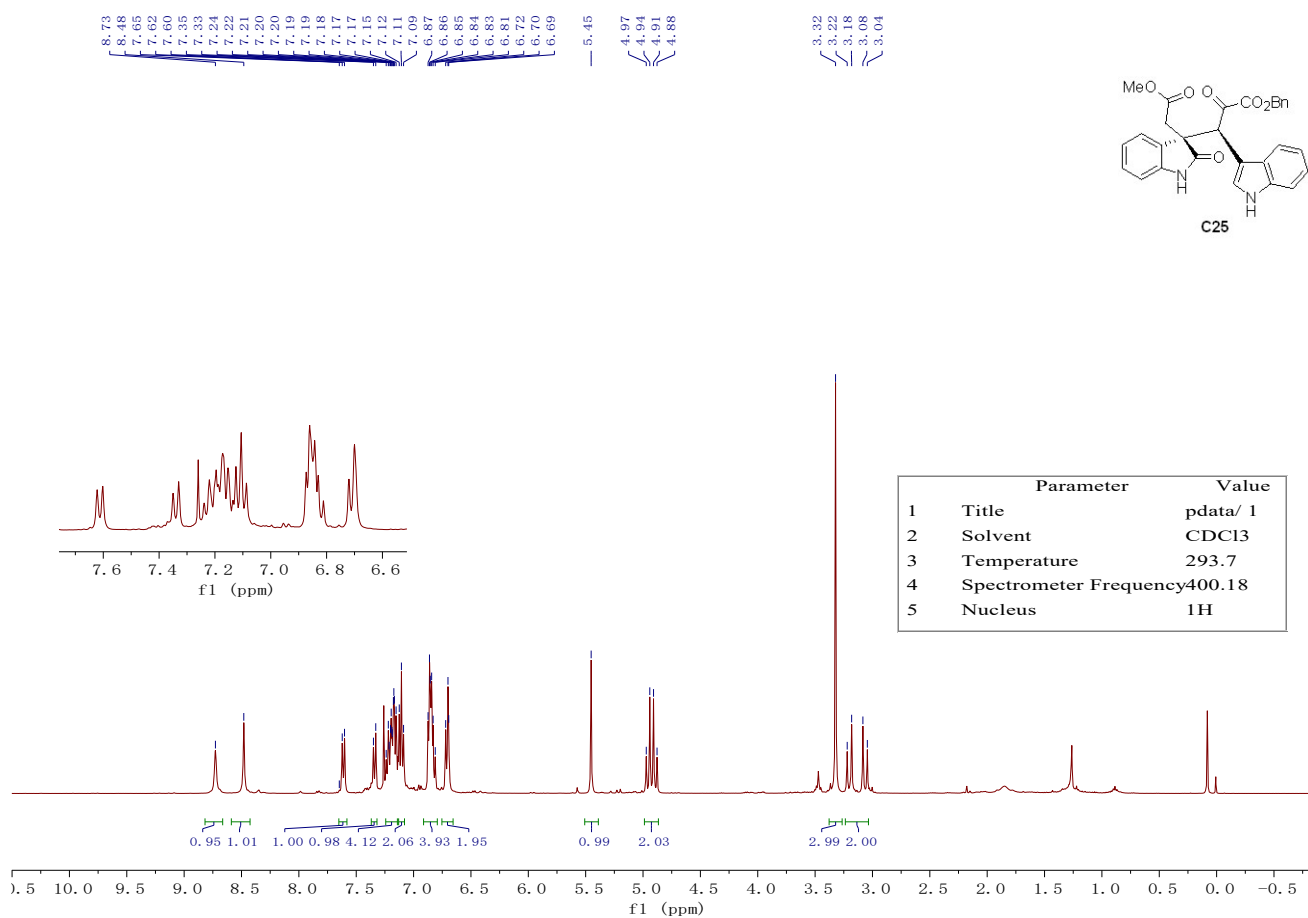
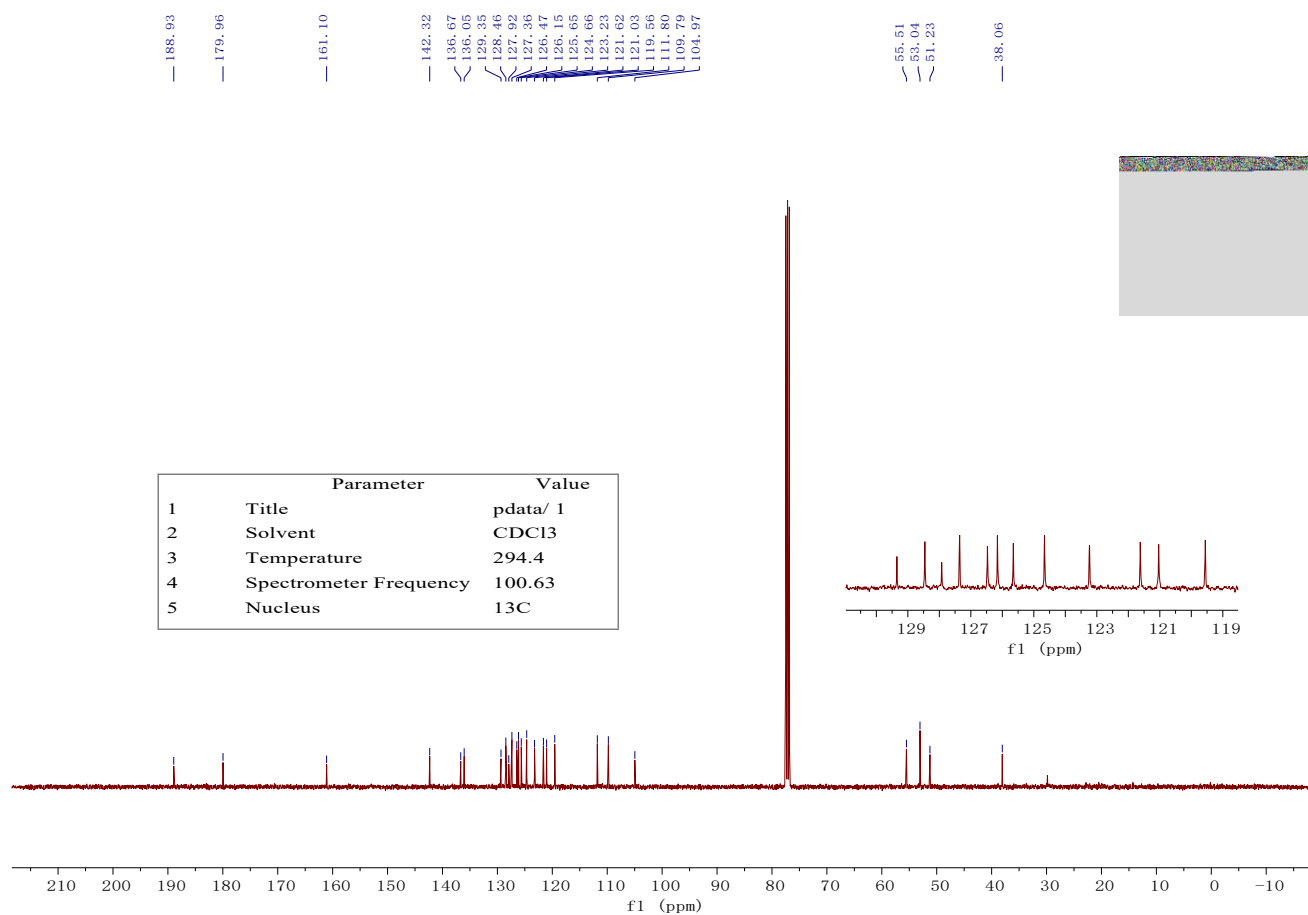


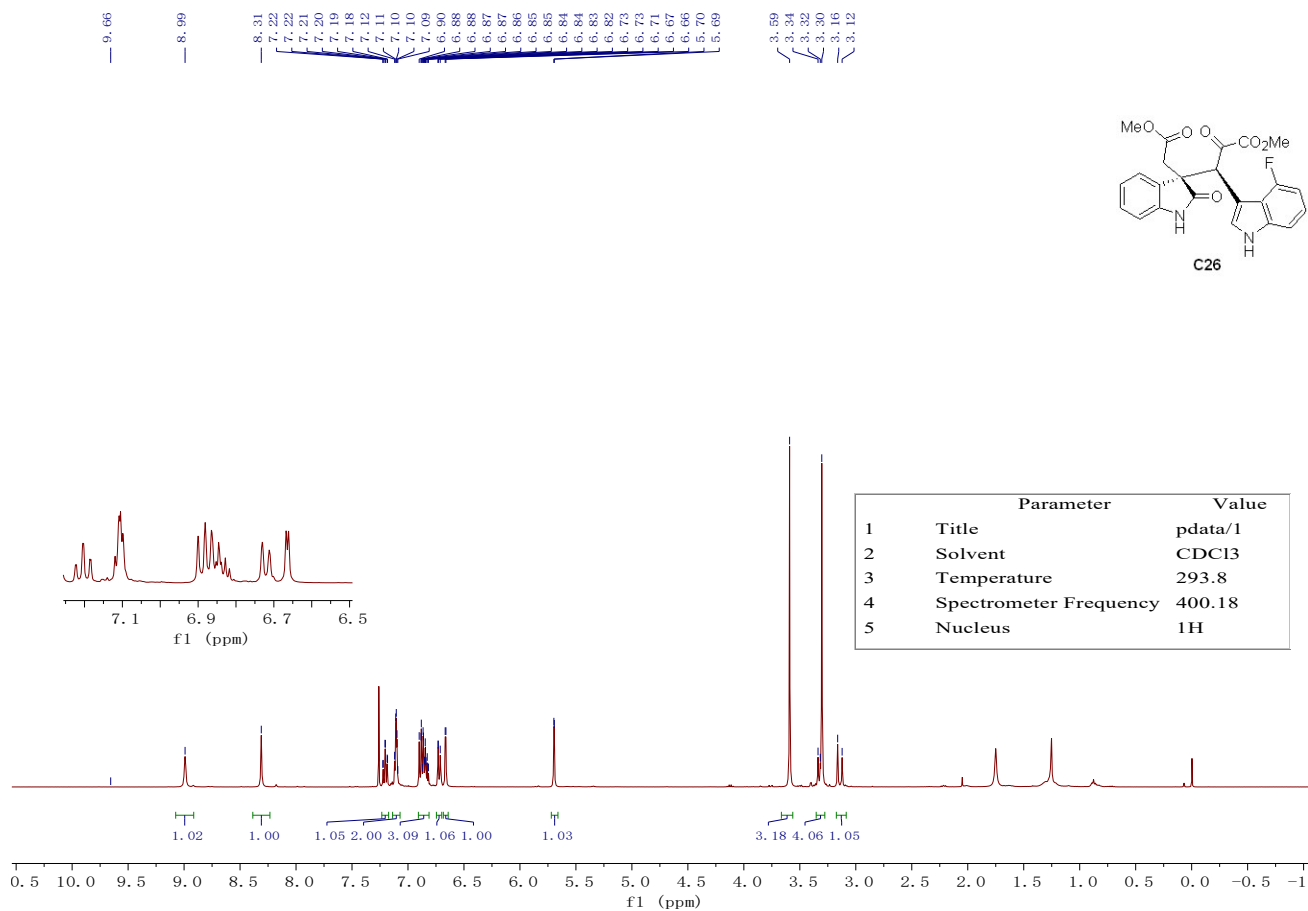
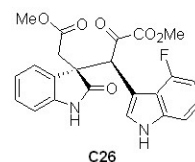
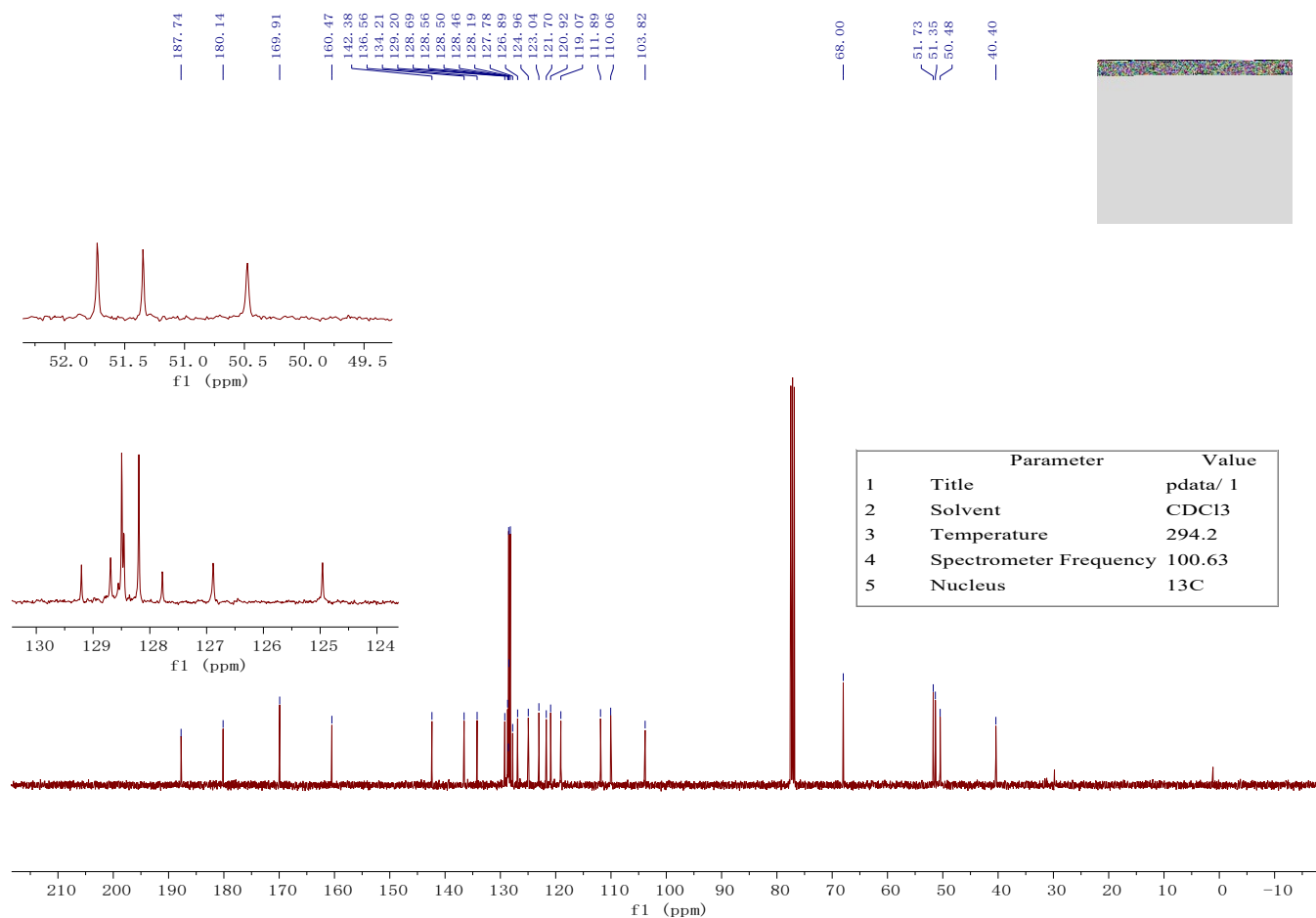


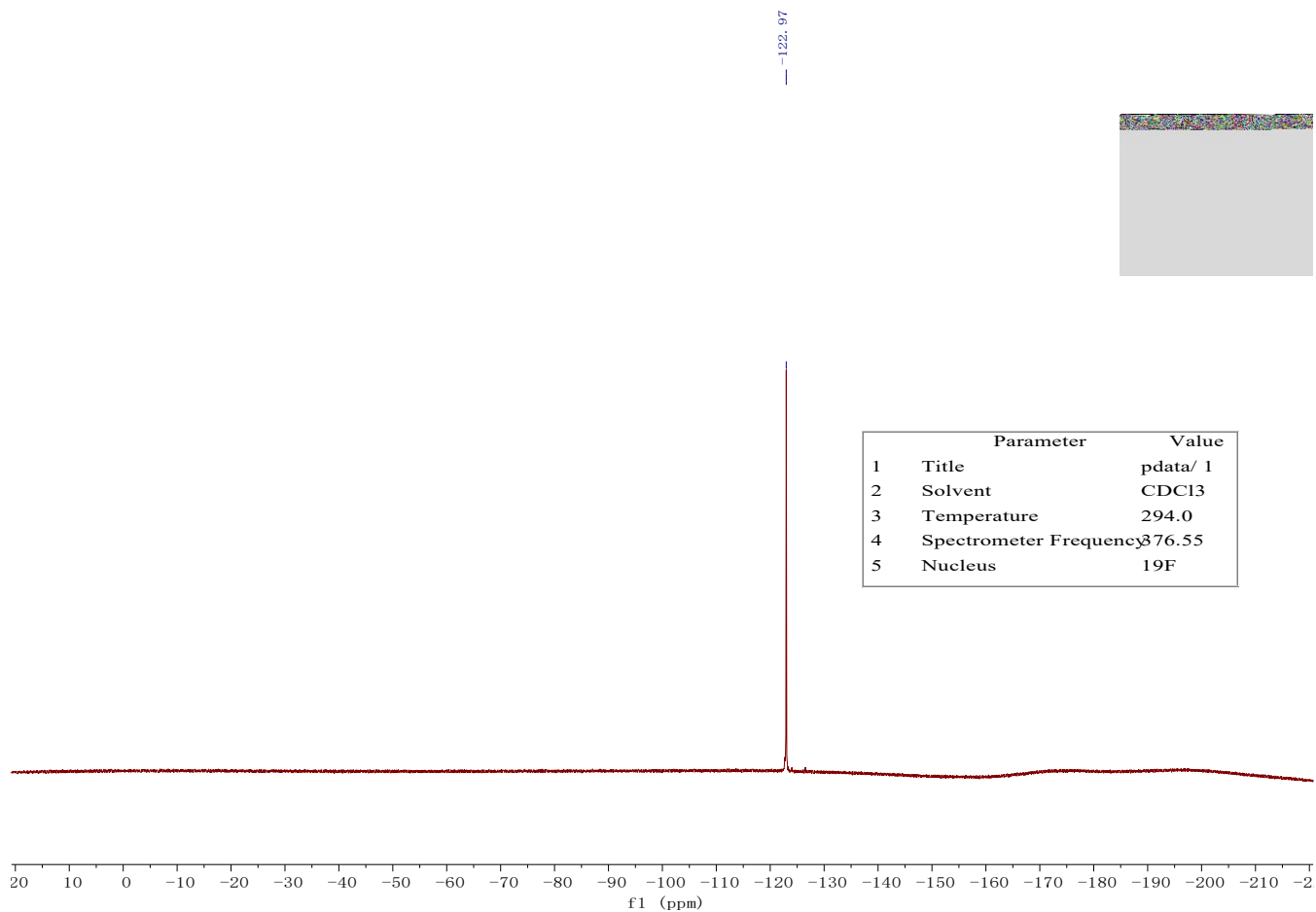
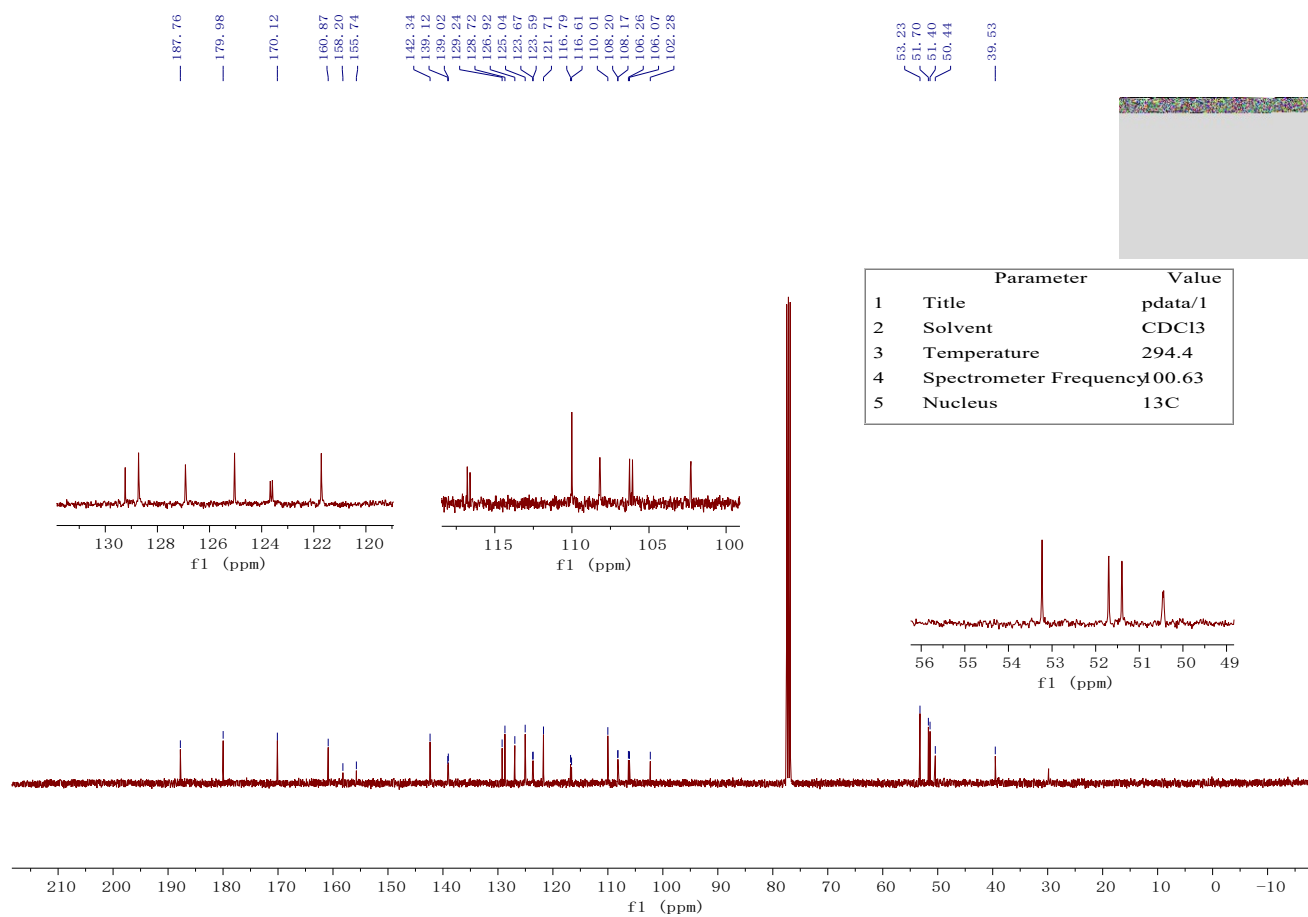


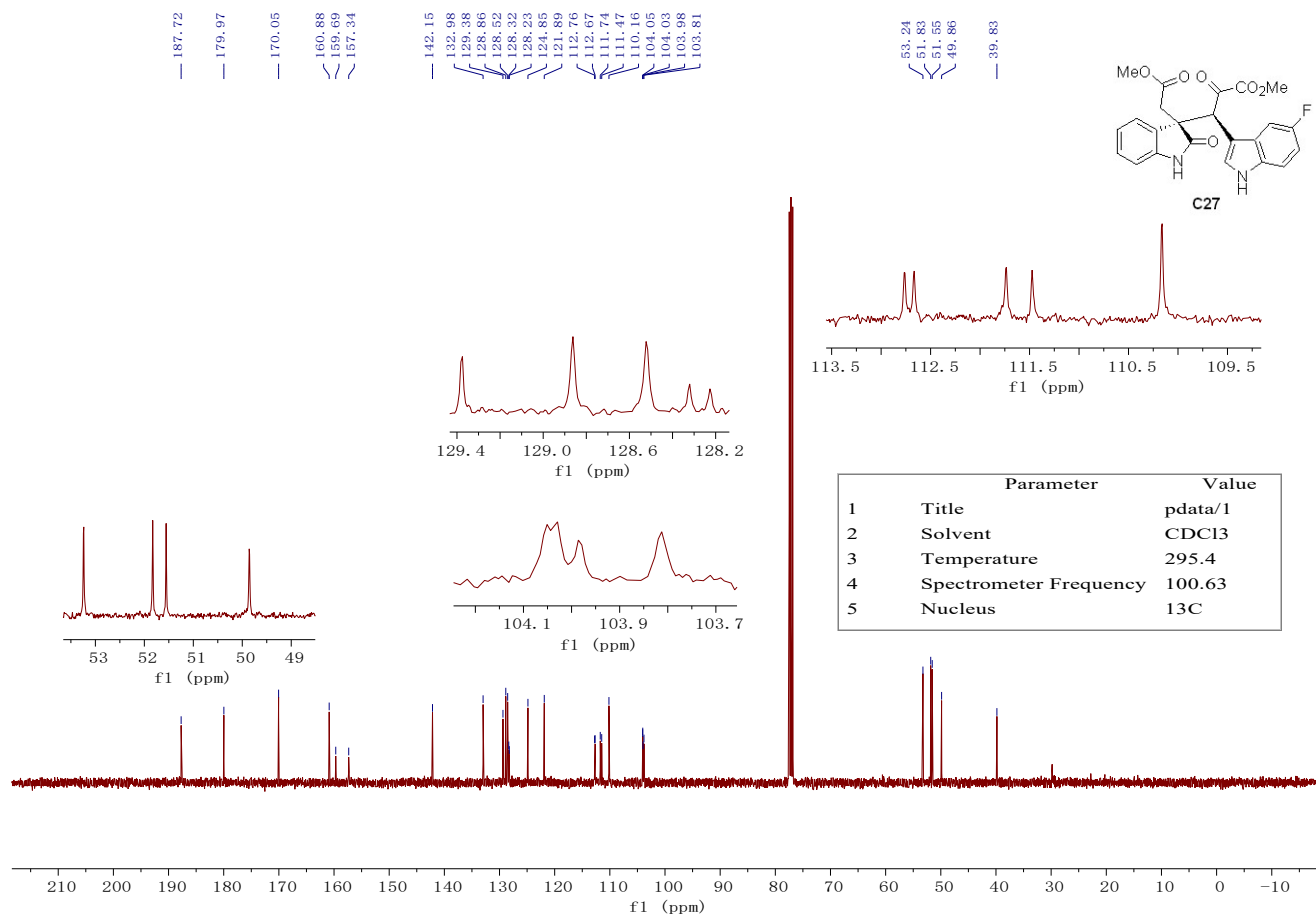
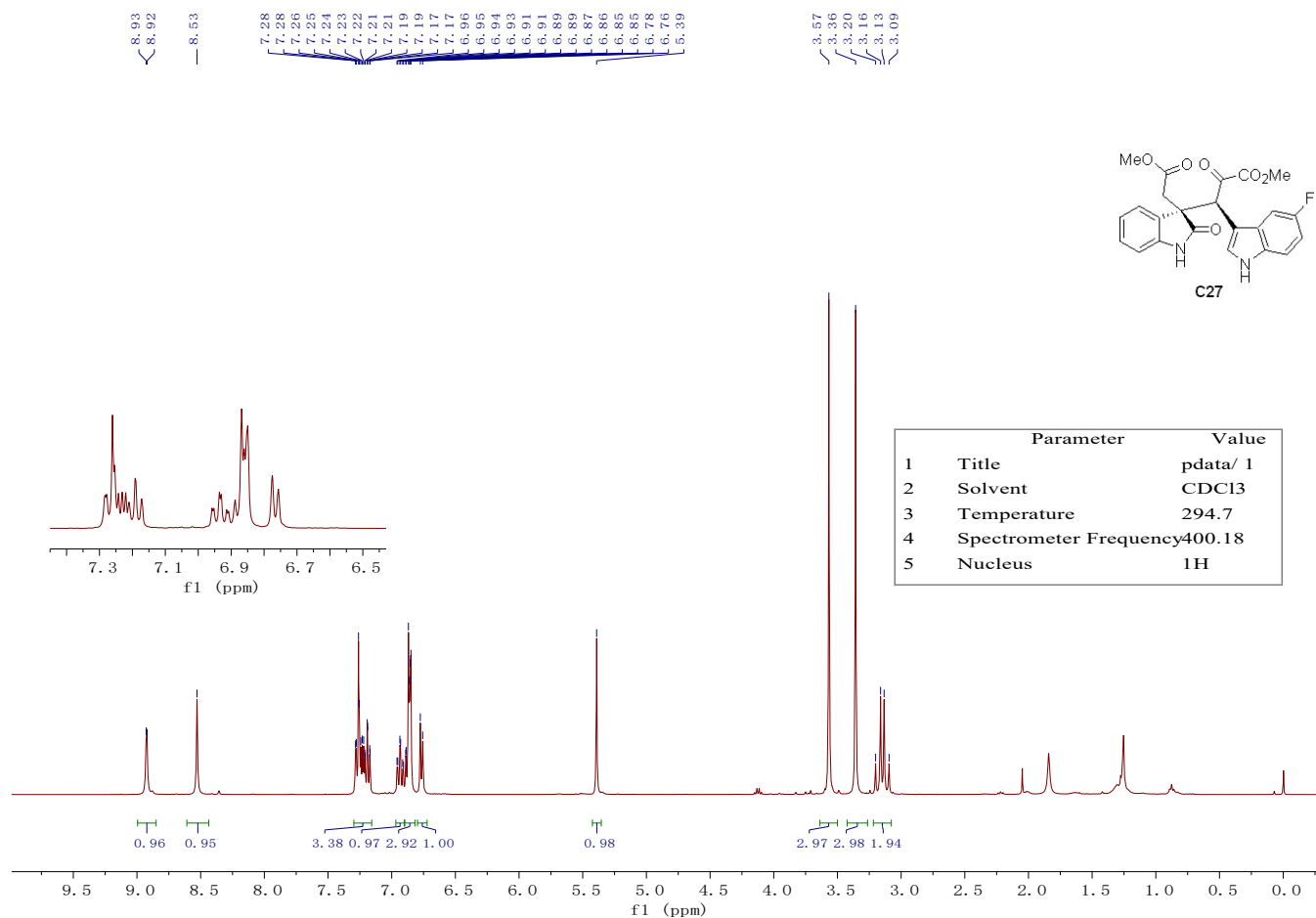
C24



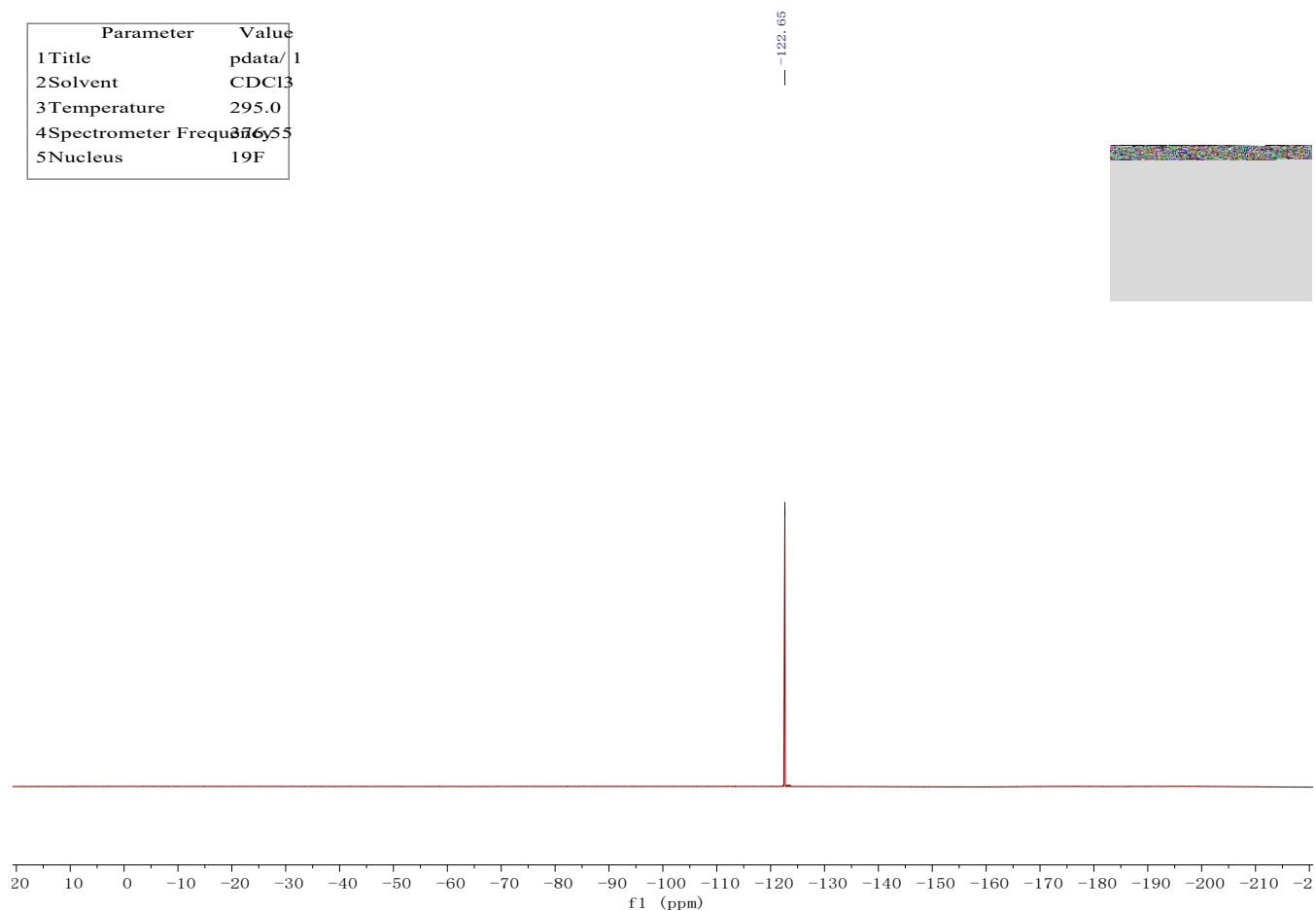




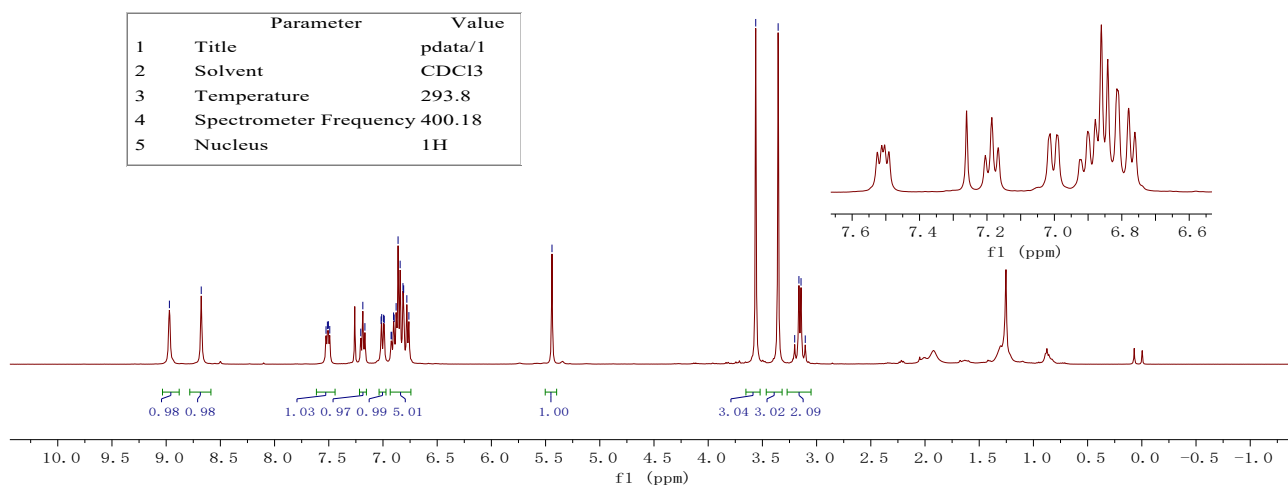
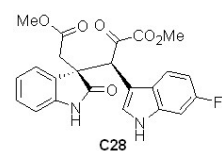




| Parameter                | Value    |
|--------------------------|----------|
| 1 Title                  | pdata/1  |
| 2 Solvent                | CDCl3    |
| 3 Temperature            | 295.0    |
| 4 Spectrometer Frequency | 500.1355 |
| 5 Nucleus                | 19F      |

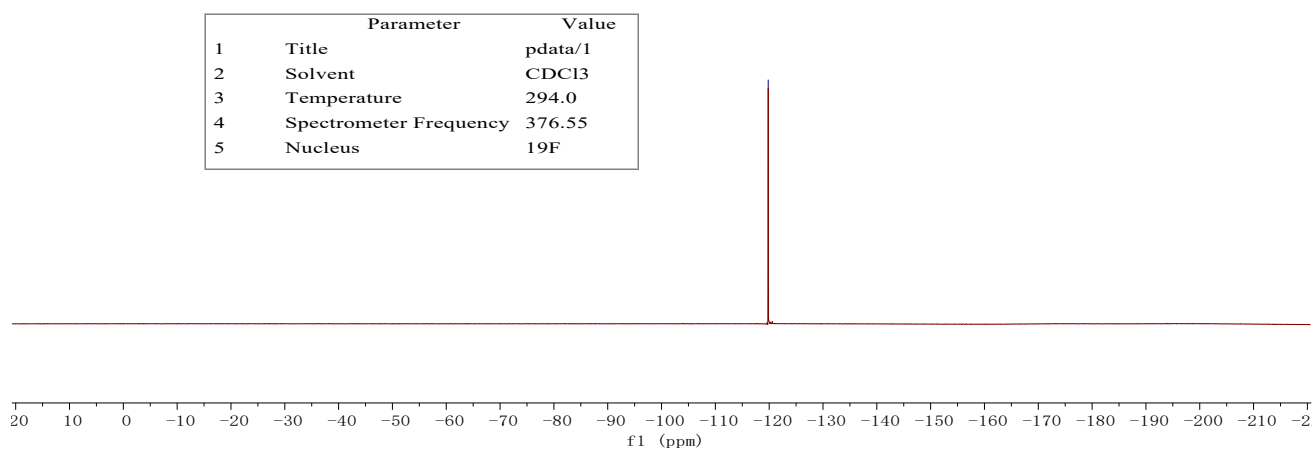
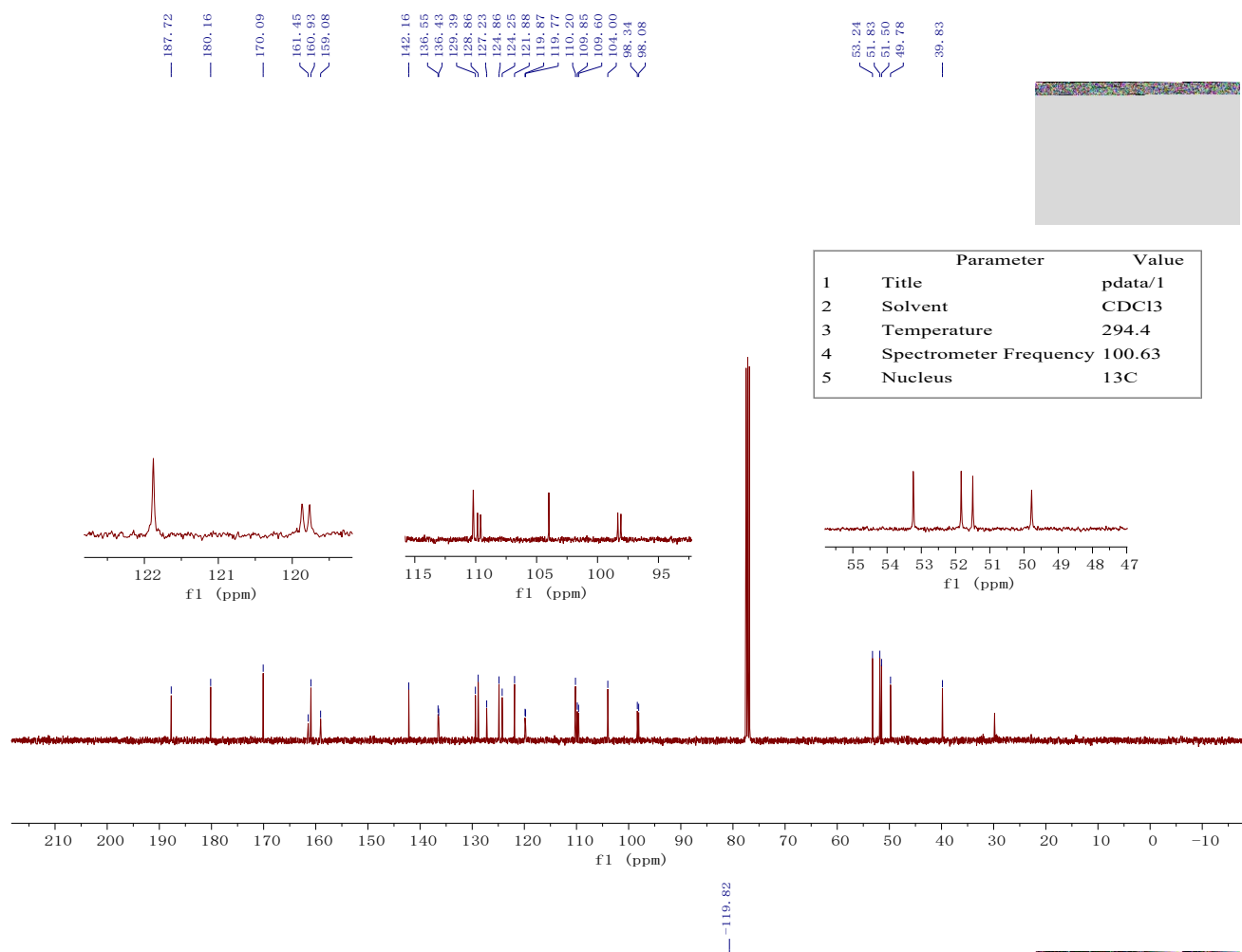


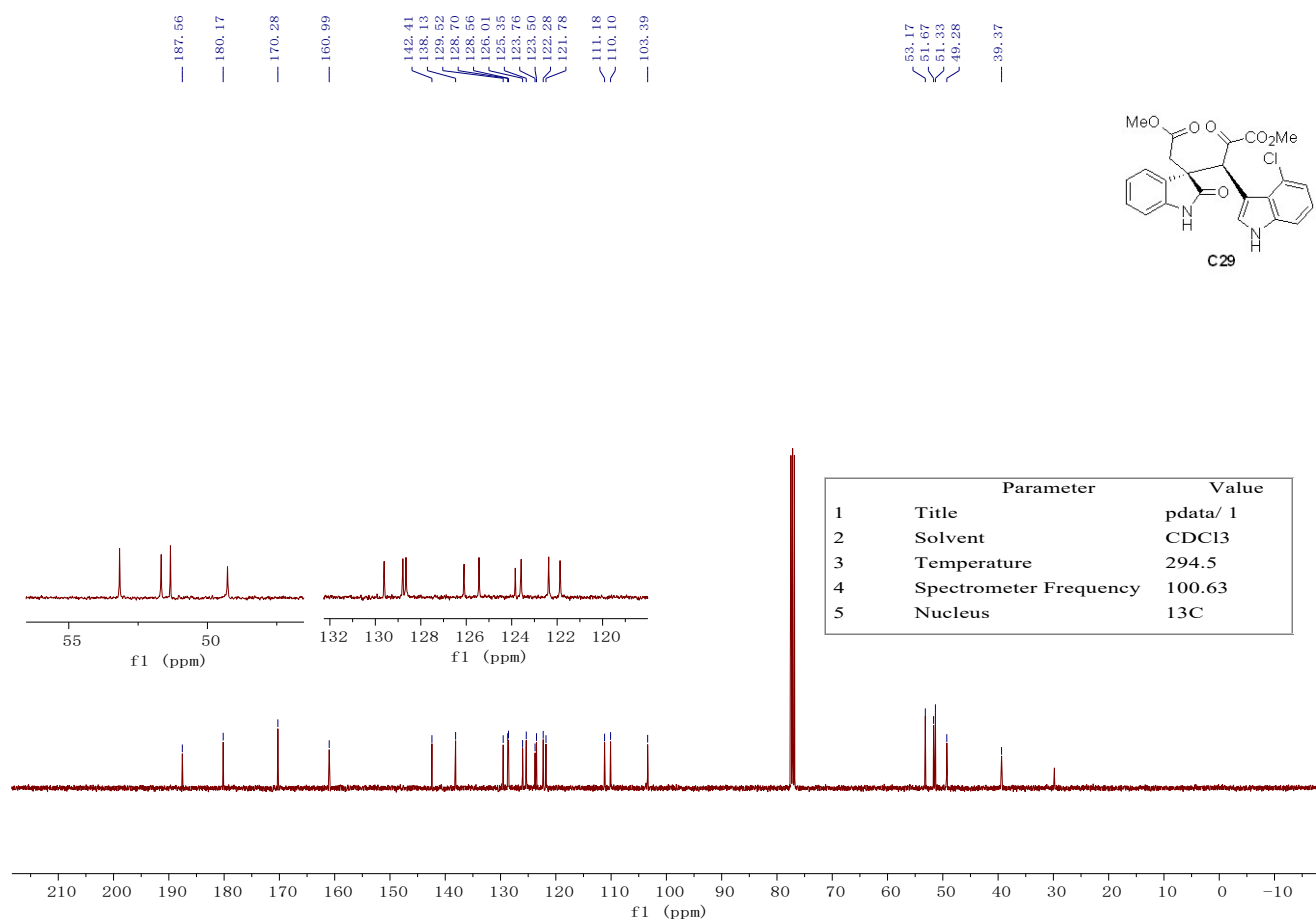
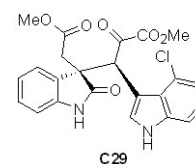
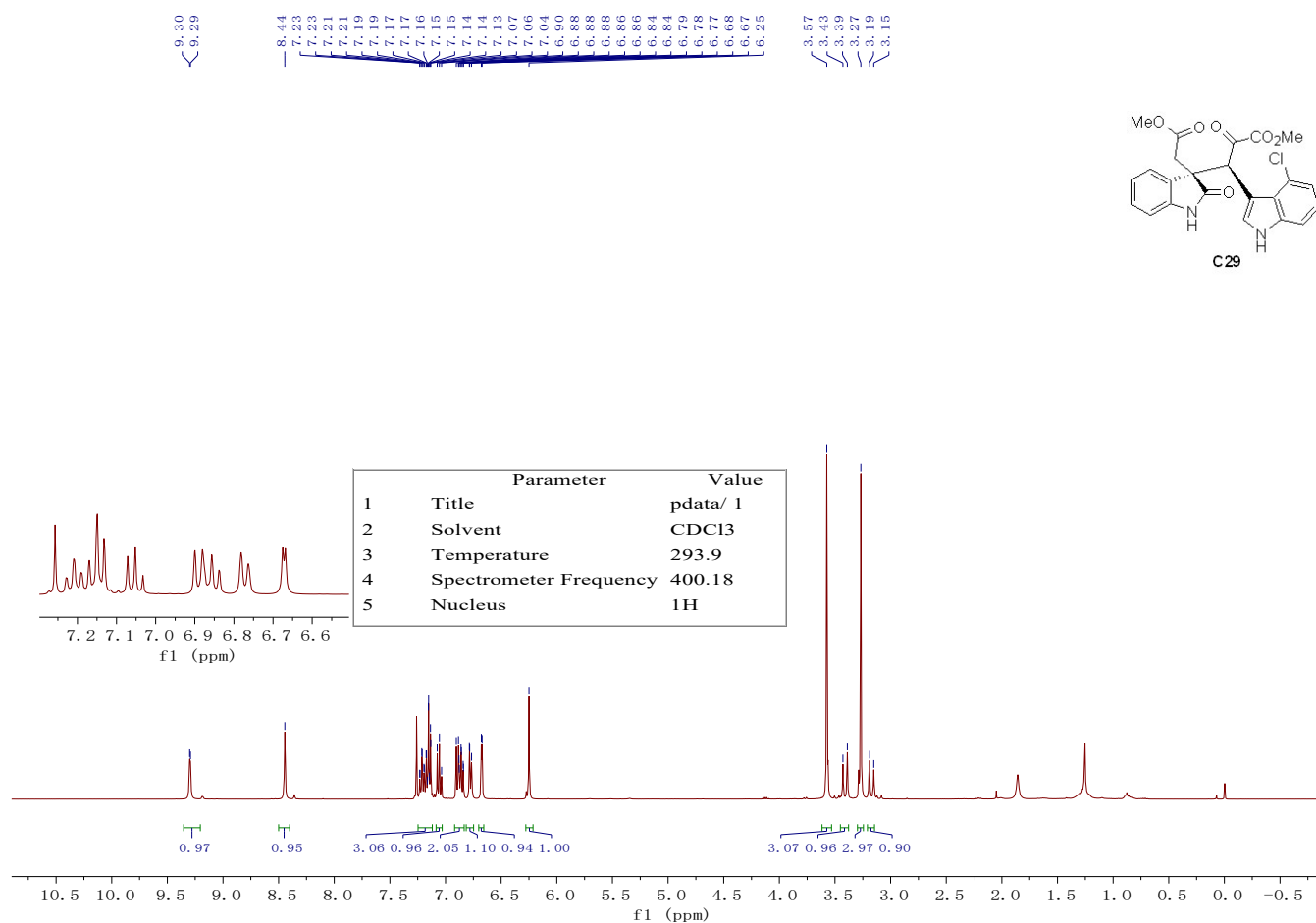
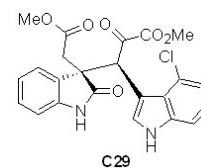
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7.50  
7.49  
7.20  
7.19  
7.17  
7.02  
6.99  
6.99  
6.93  
6.92  
6.90  
6.90  
6.88  
6.86  
6.84  
6.82  
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3.14  
3.10

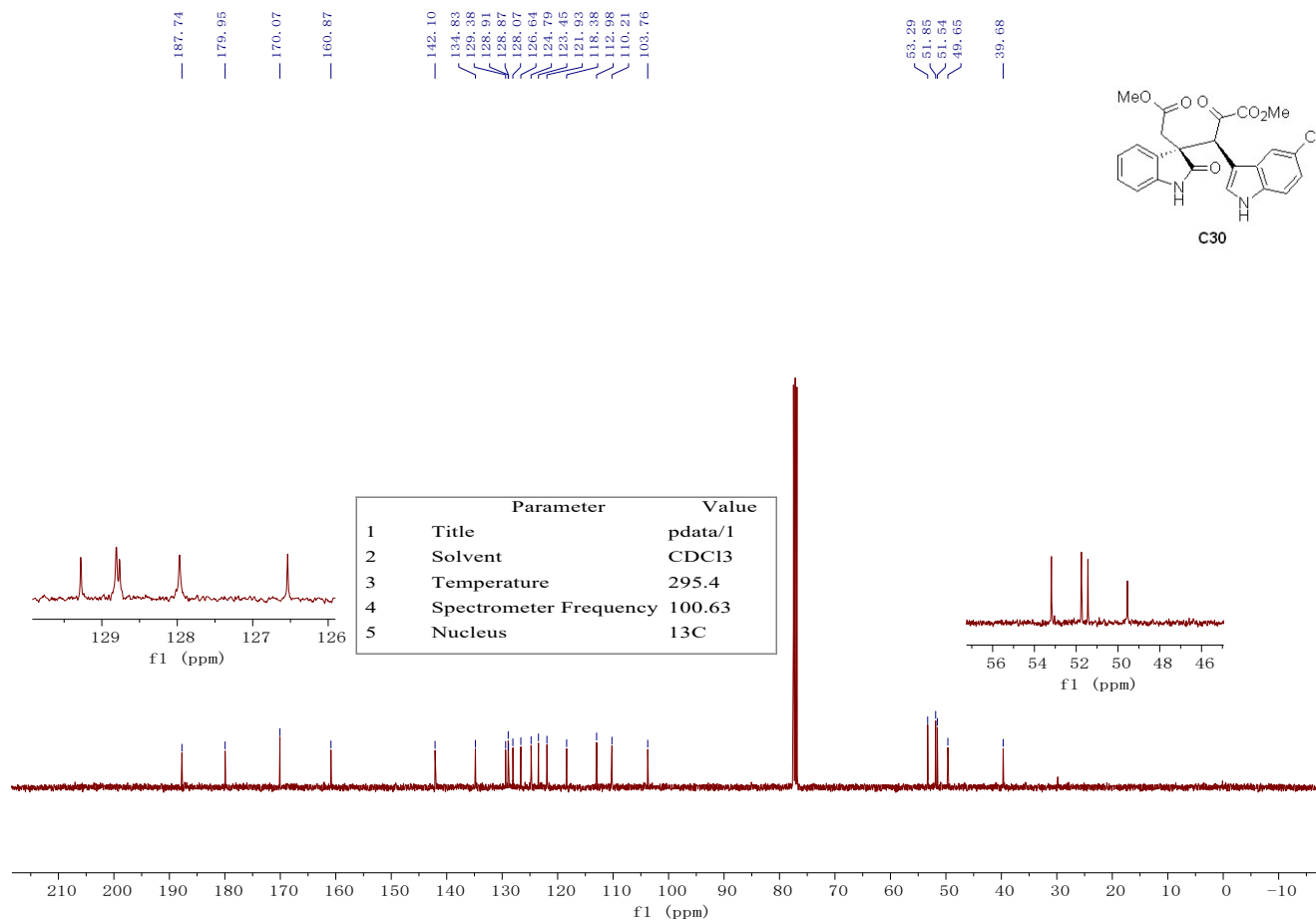
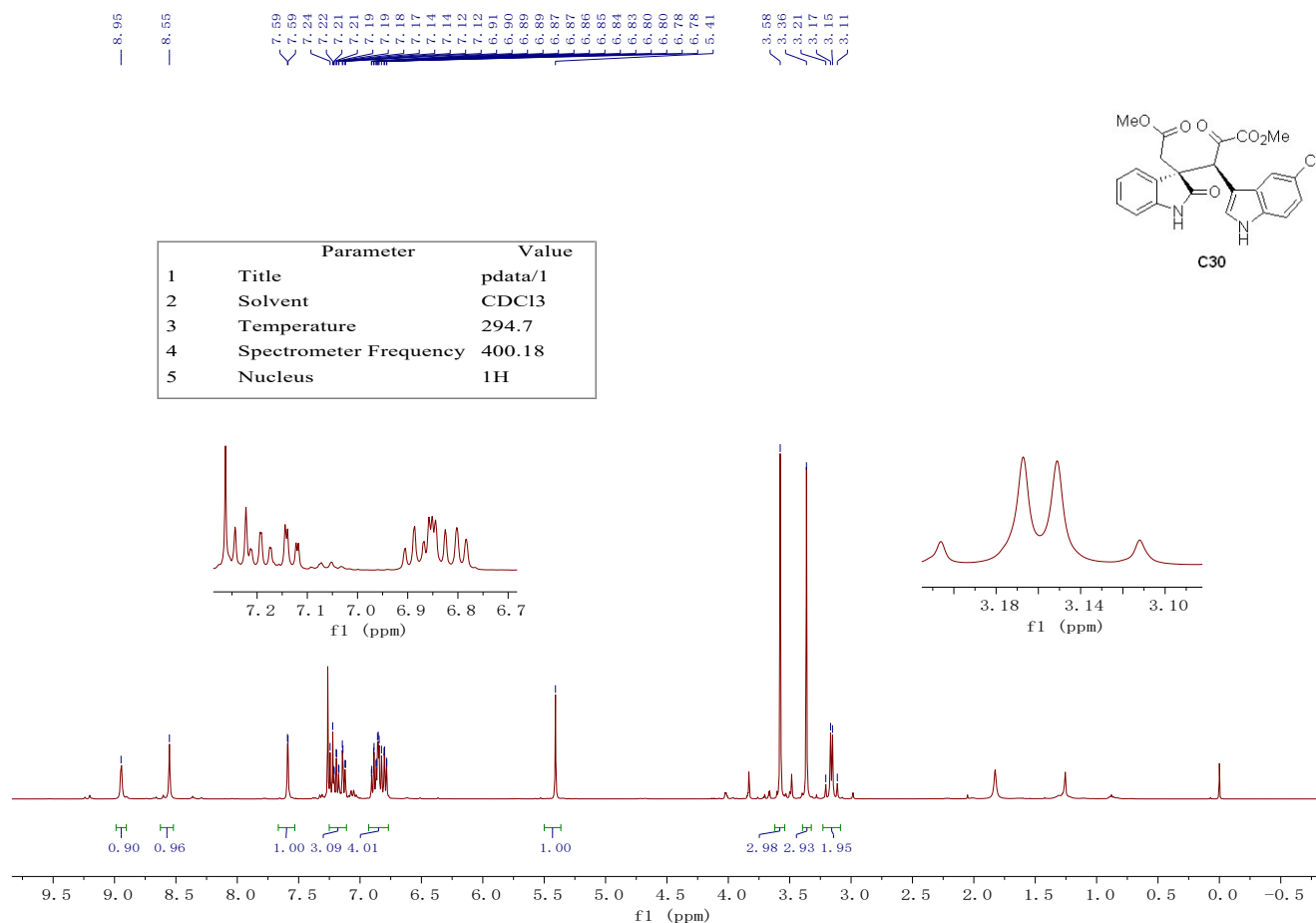


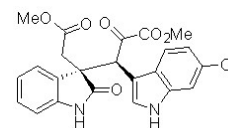
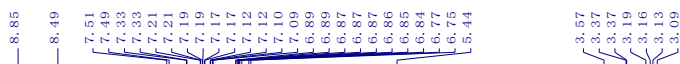
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| 1 Title                  | pdata/1 |
| 2 Solvent                | CDCl3   |
| 3 Temperature            | 293.8   |
| 4 Spectrometer Frequency | 400.18  |
| 5 Nucleus                | 1H      |

0.98 0.98  
1.03 0.97 0.99 5.01  
1.00  
3.04 3.02 2.09

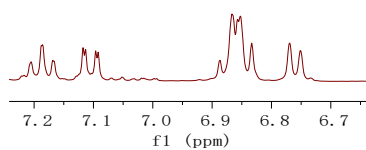




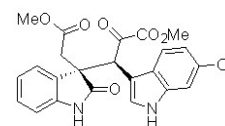
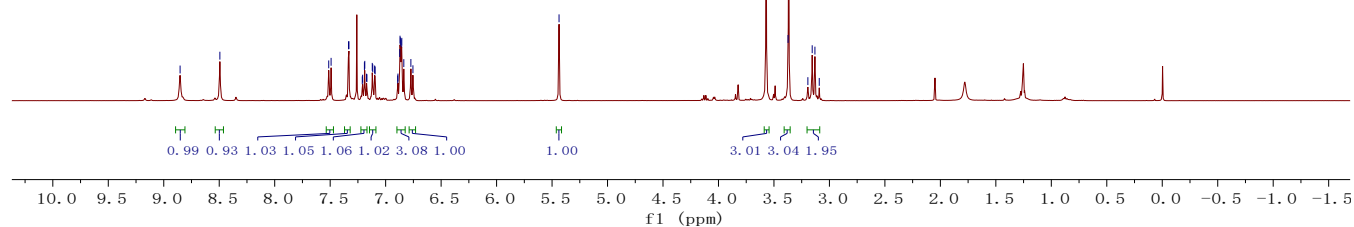




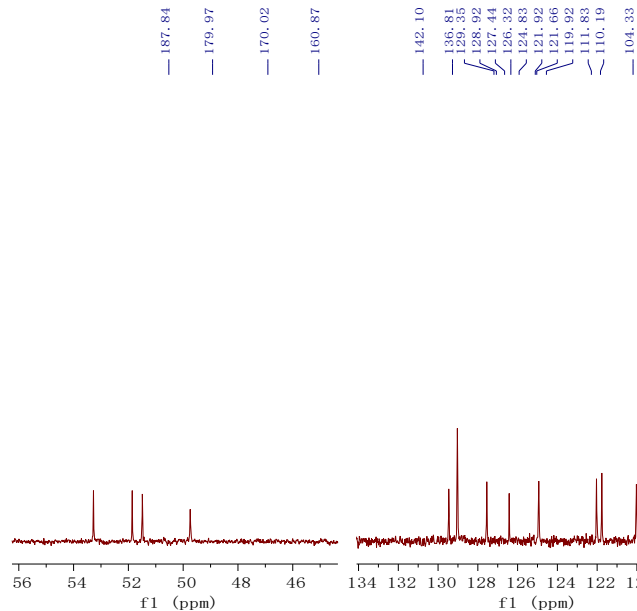
C31



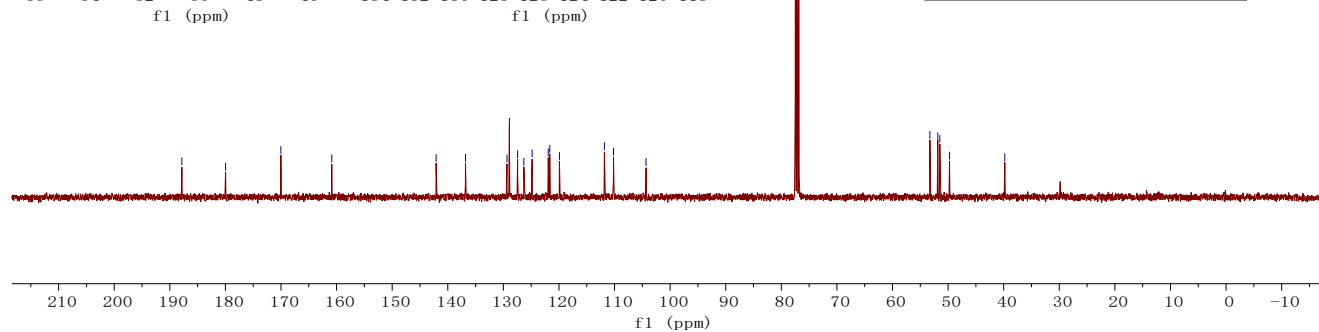
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|--------------------------|----------------|
| 1 Title                  | pdata/ 1       |
| 2 Solvent                | CDC13          |
| 3 Temperature            | 294.7          |
| 4 Spectrometer Frequency | 400.18         |
| 5 Nucleus                | <sup>1</sup> H |

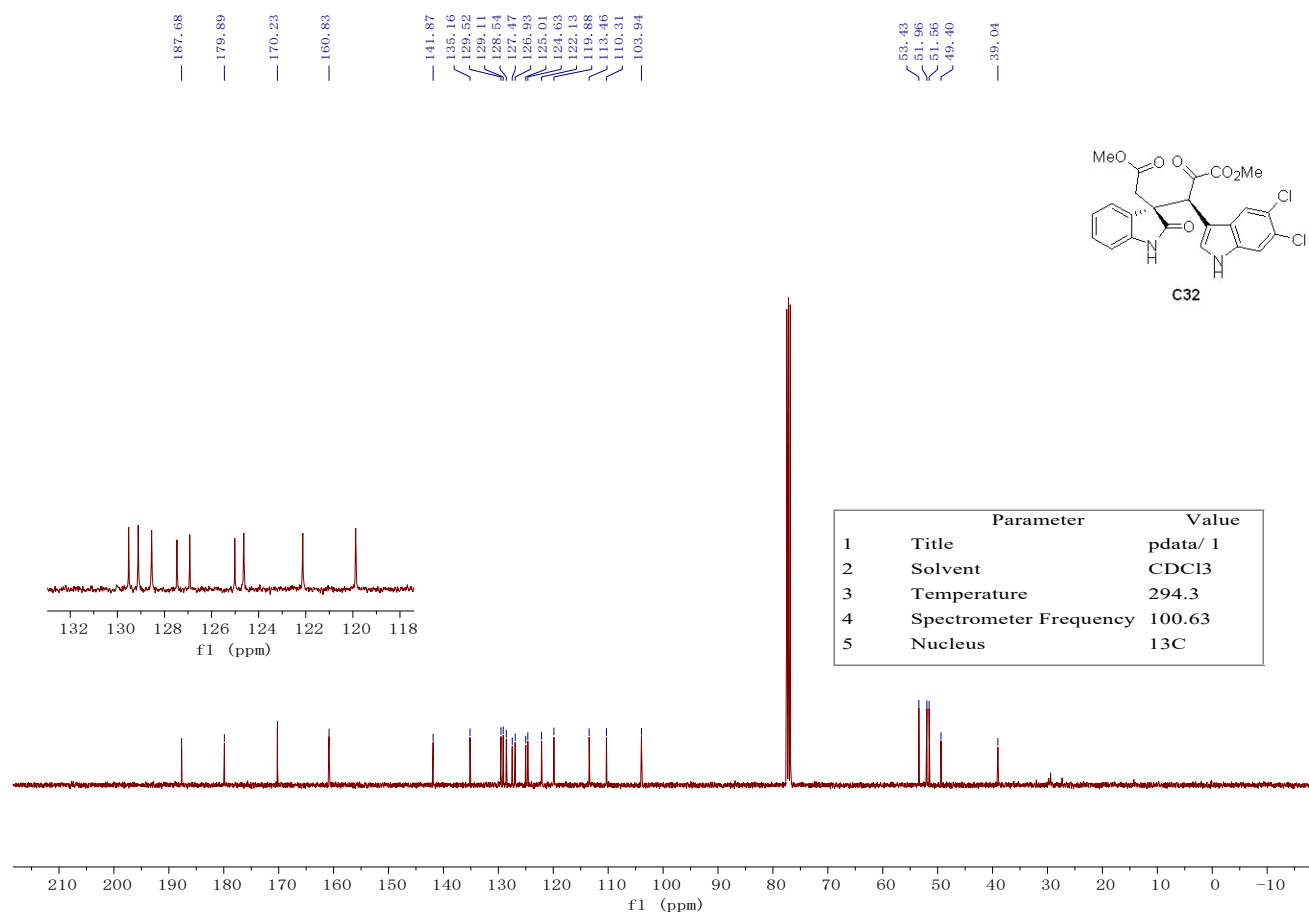
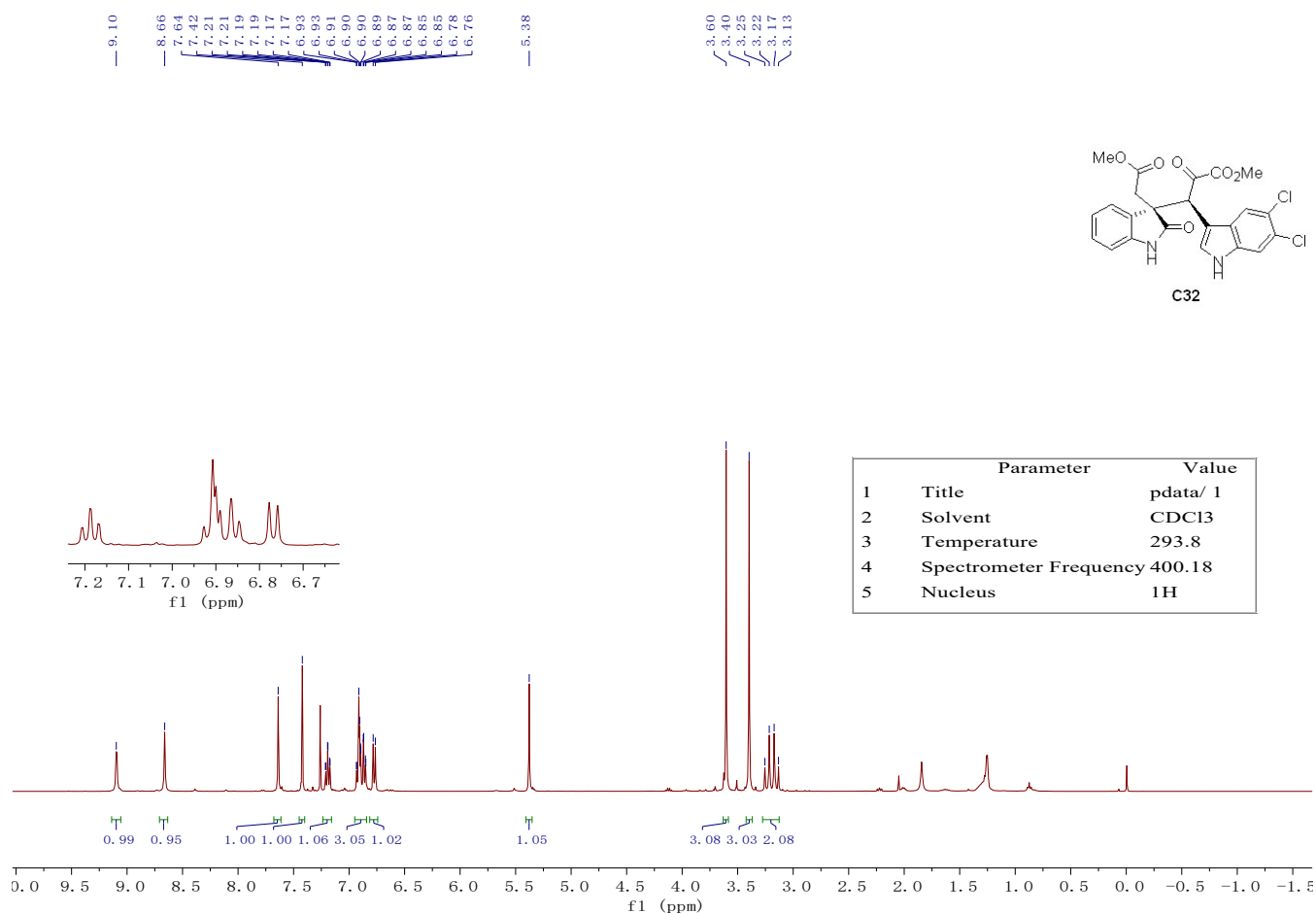


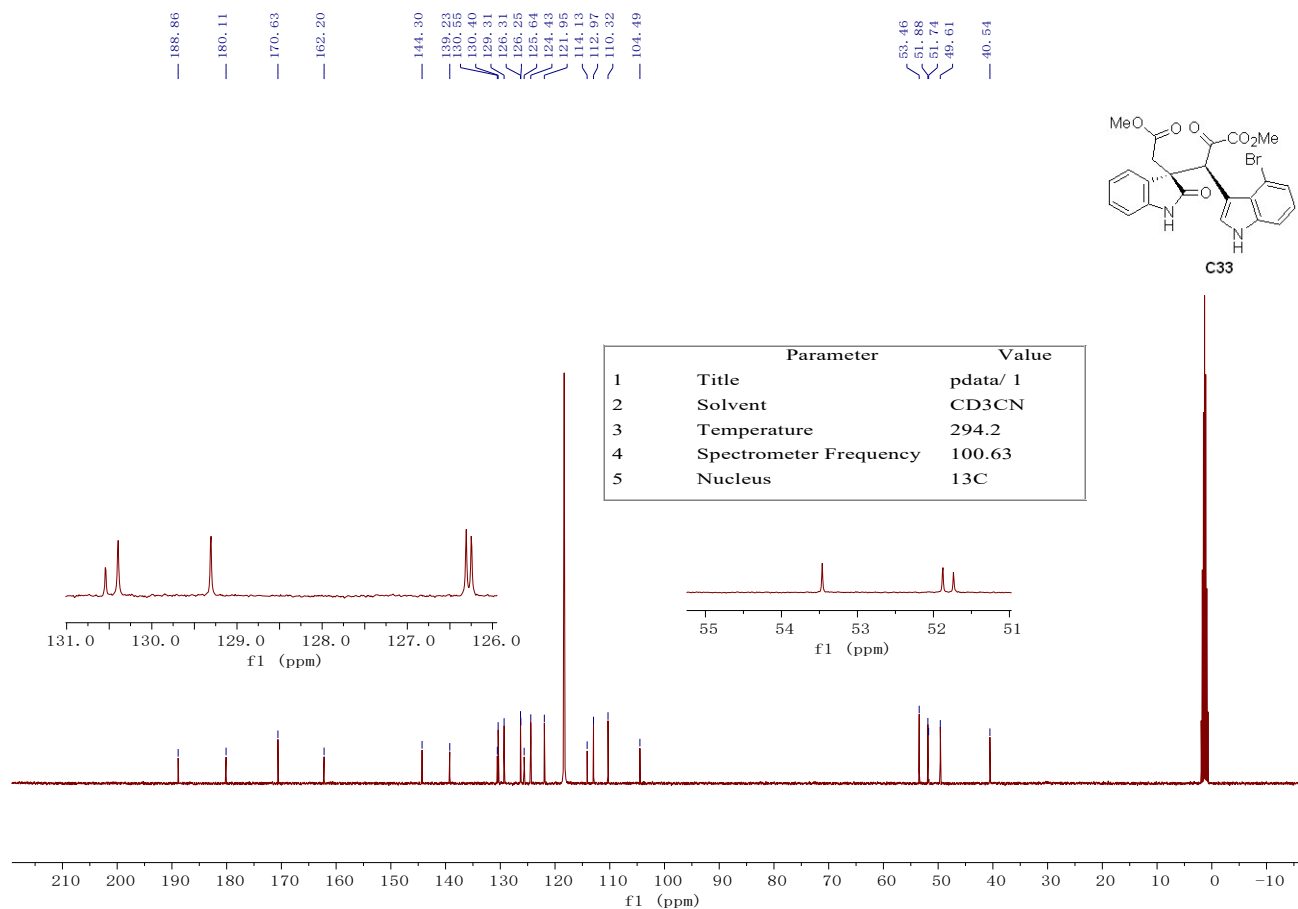
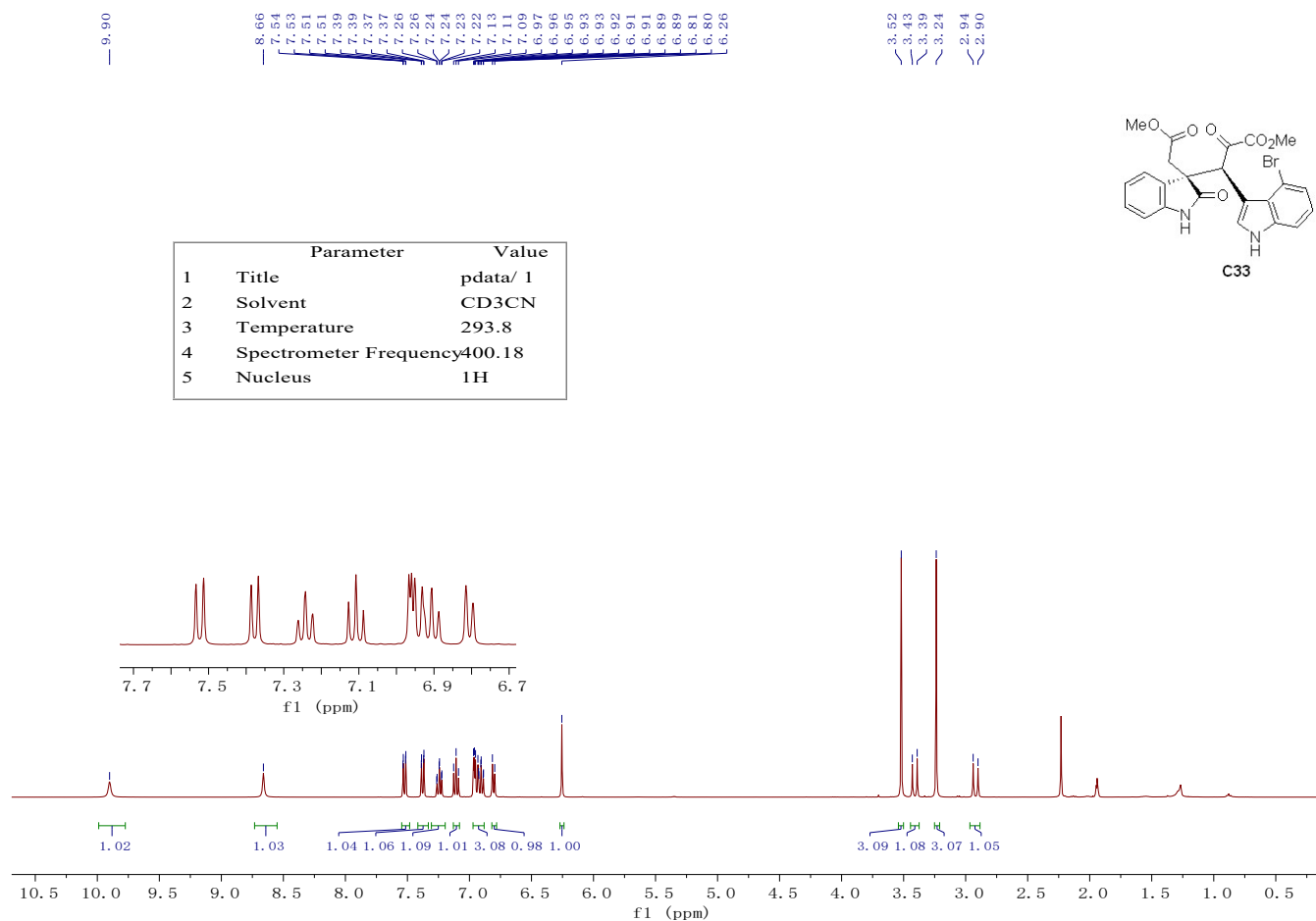
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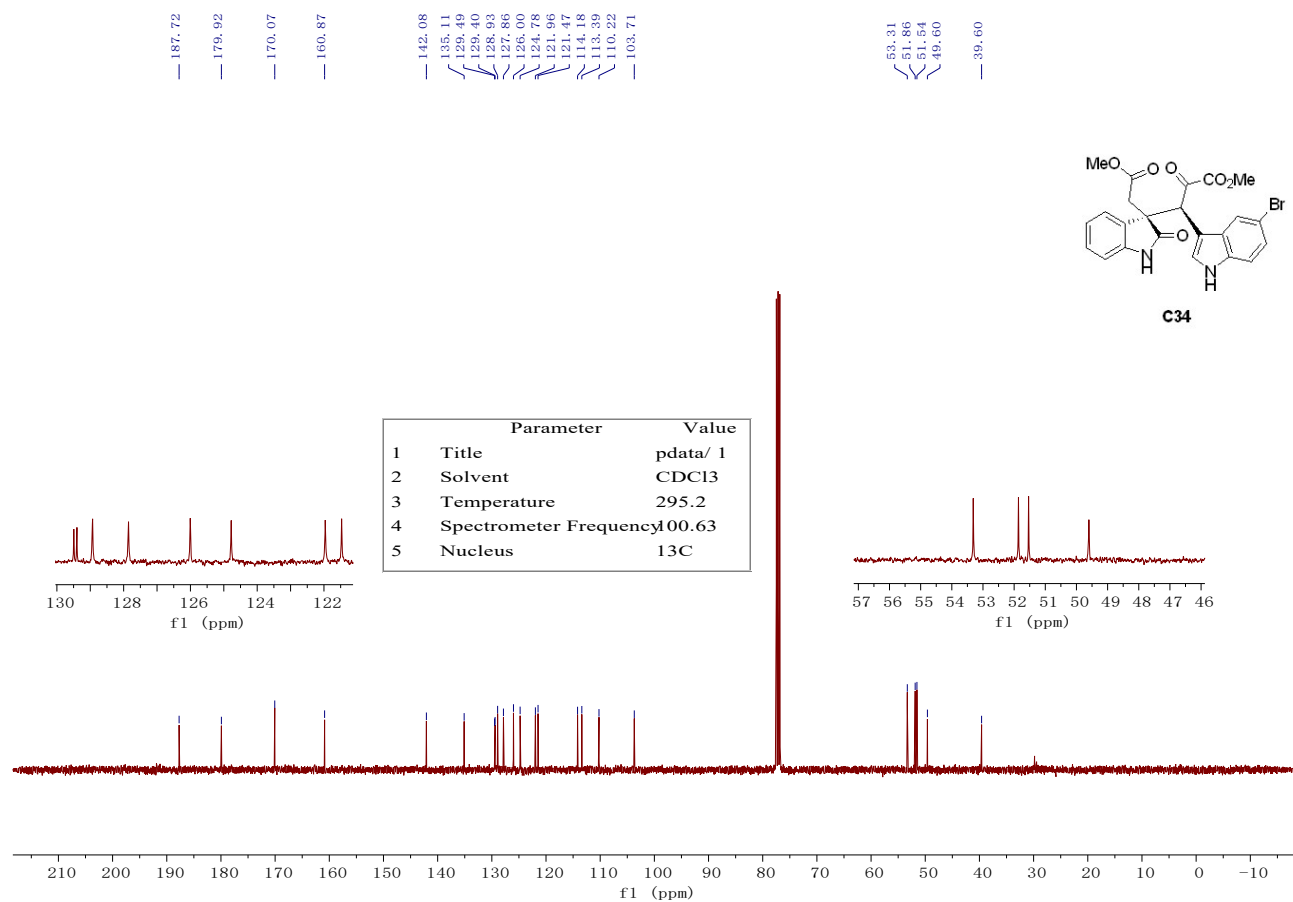
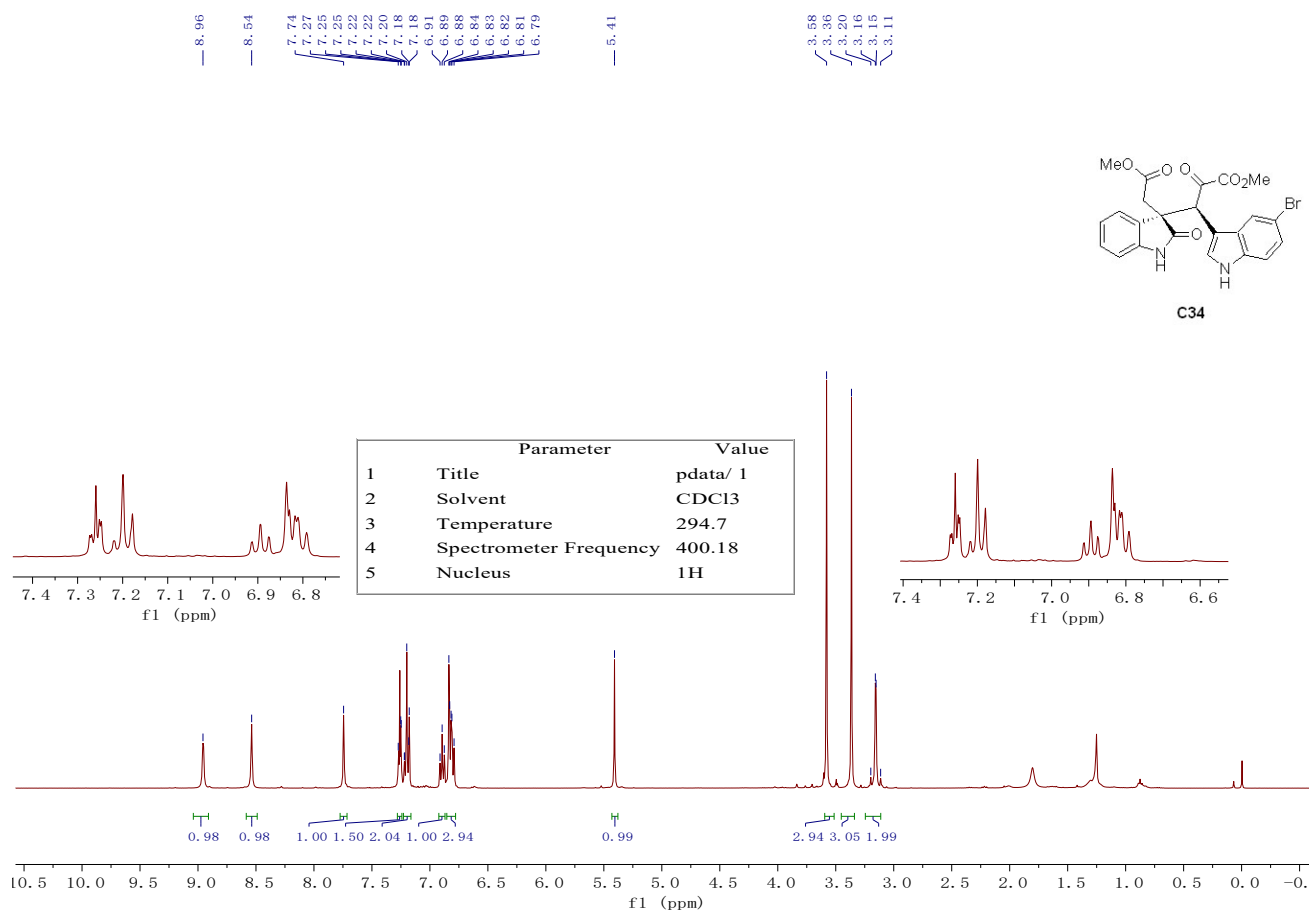


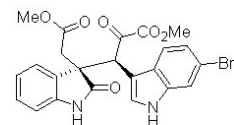
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|--------------------------|-----------------|
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| 2 Solvent                | CDC13           |
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| 4 Spectrometer Frequency | 100.63          |
| 5 Nucleus                | <sup>13</sup> C |



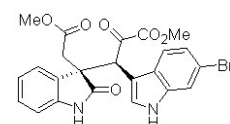
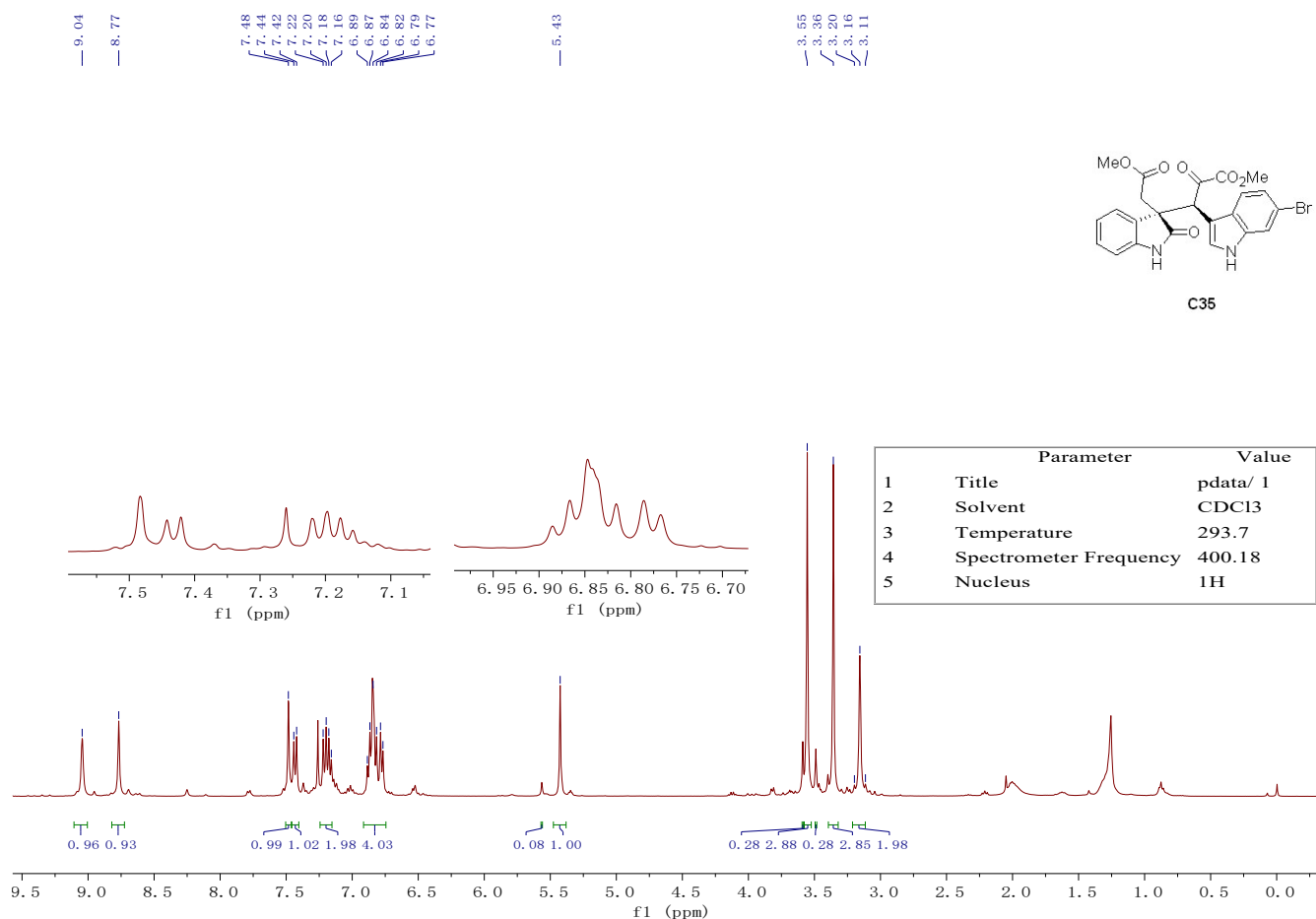




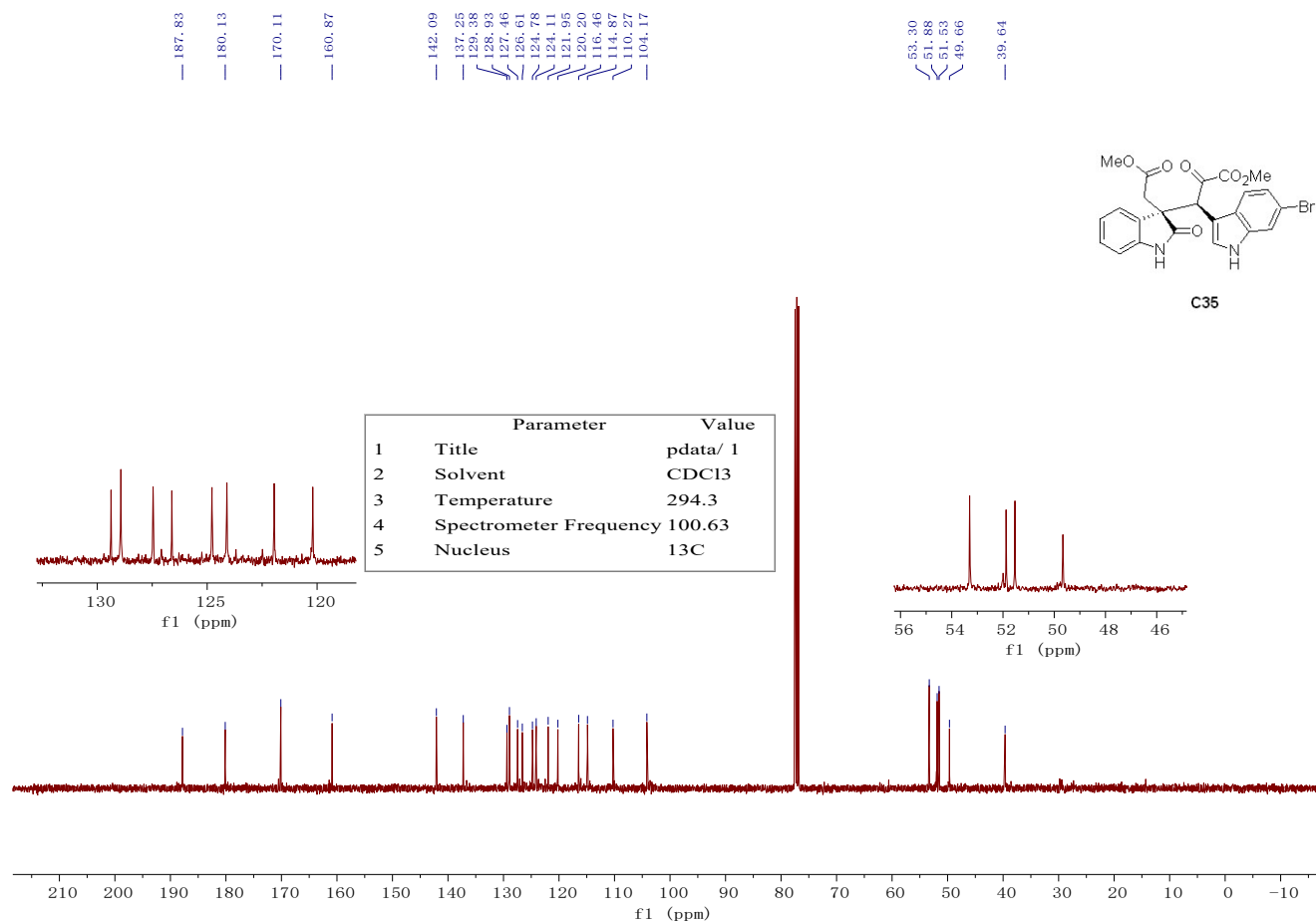


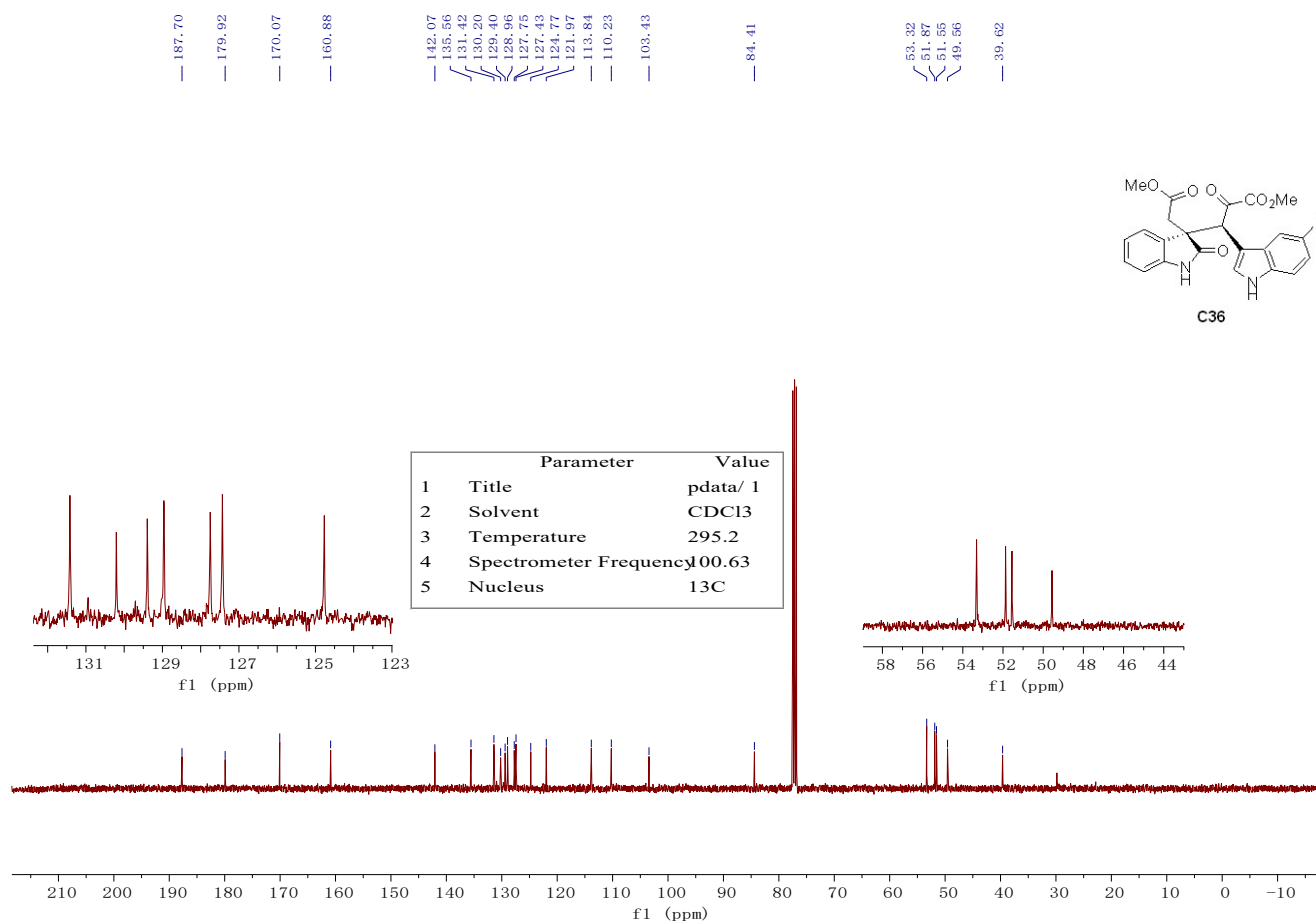
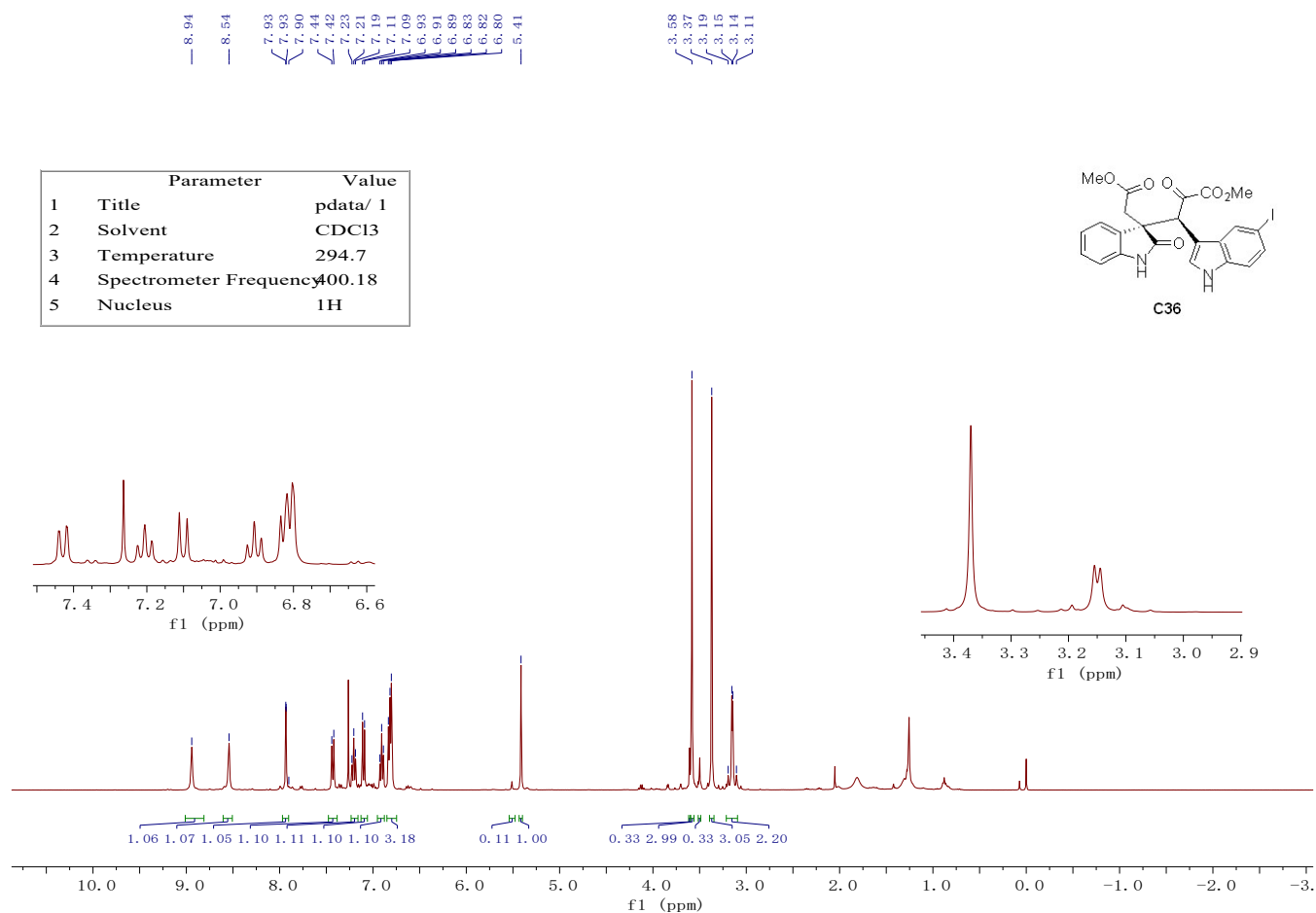


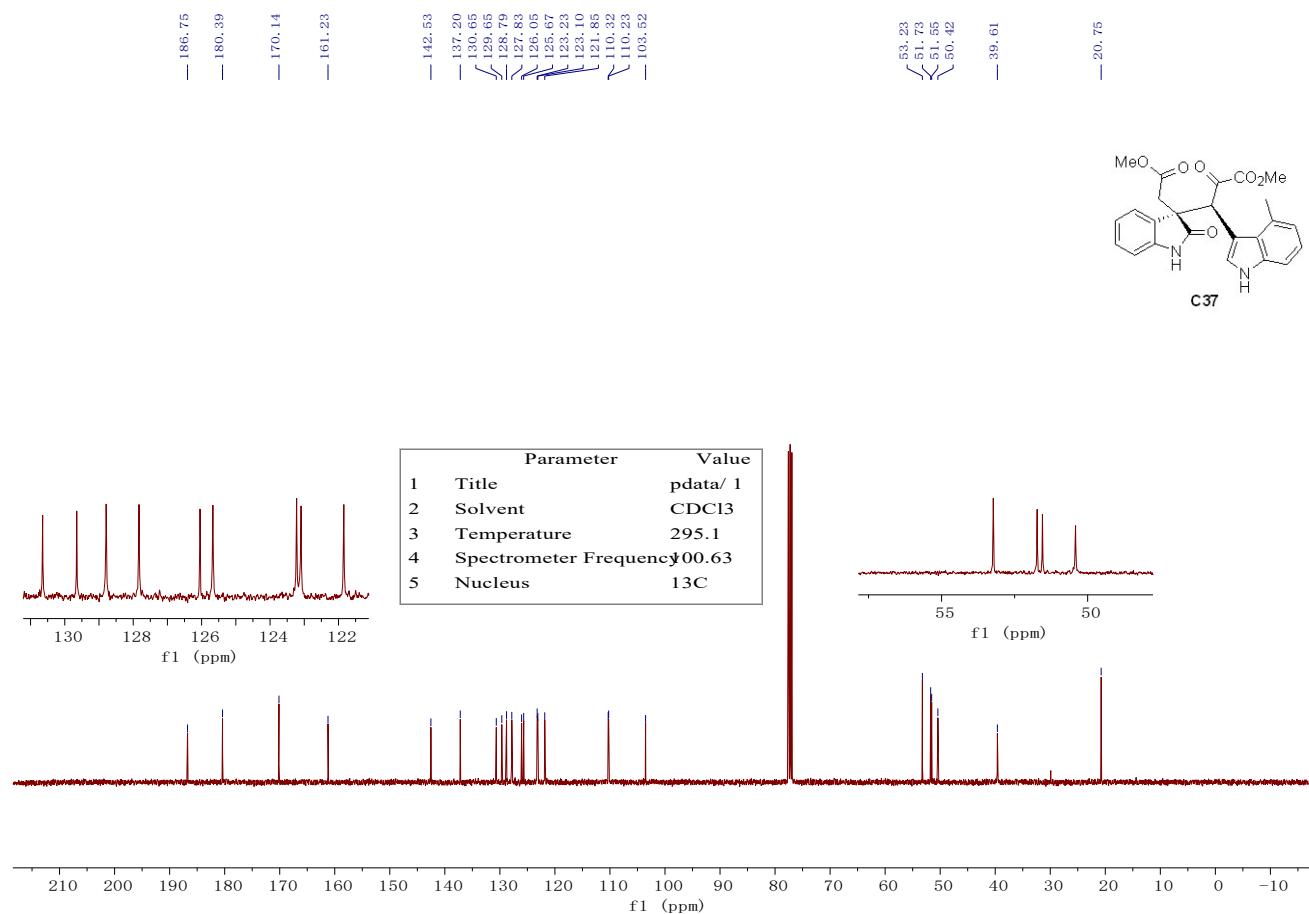
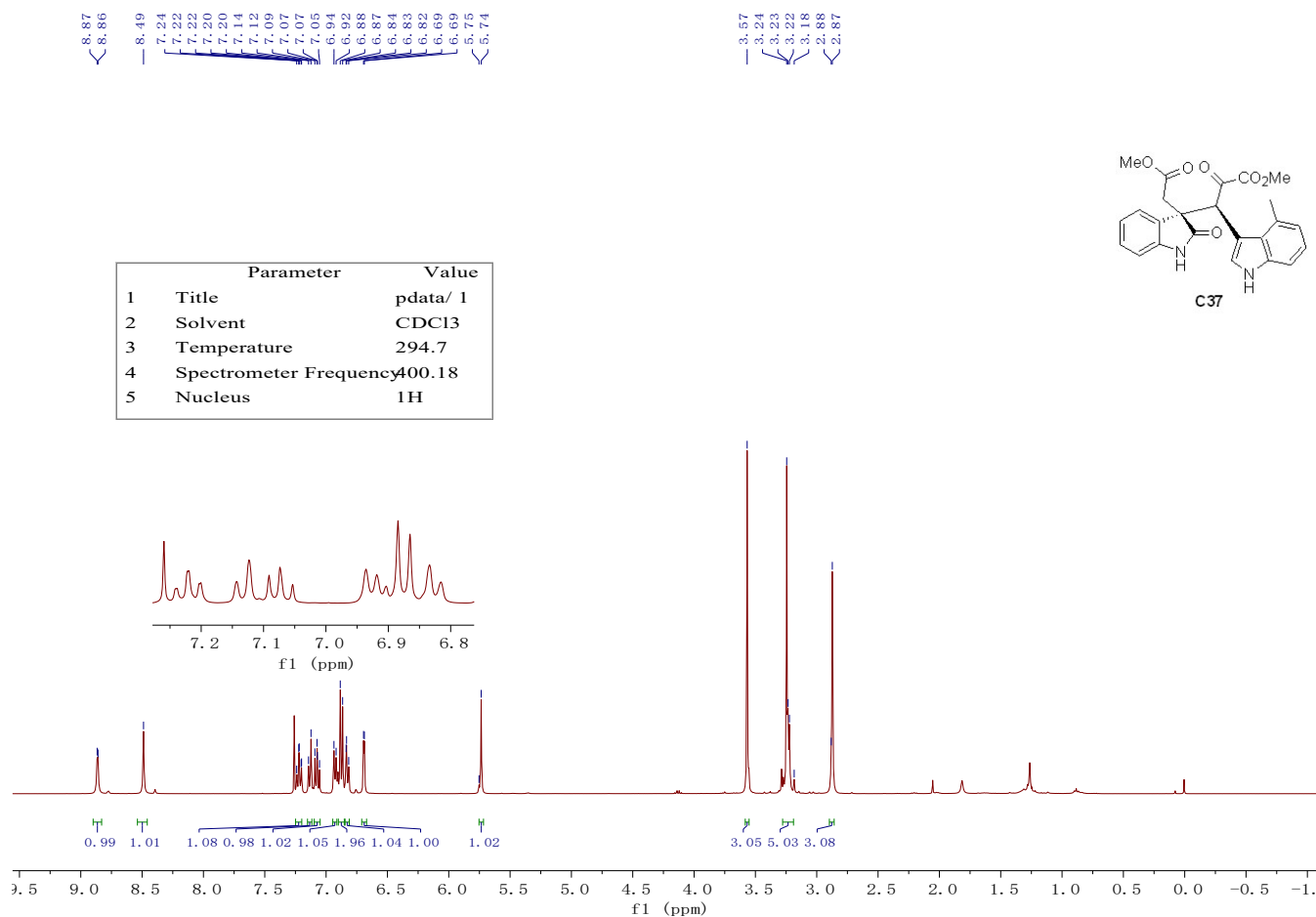
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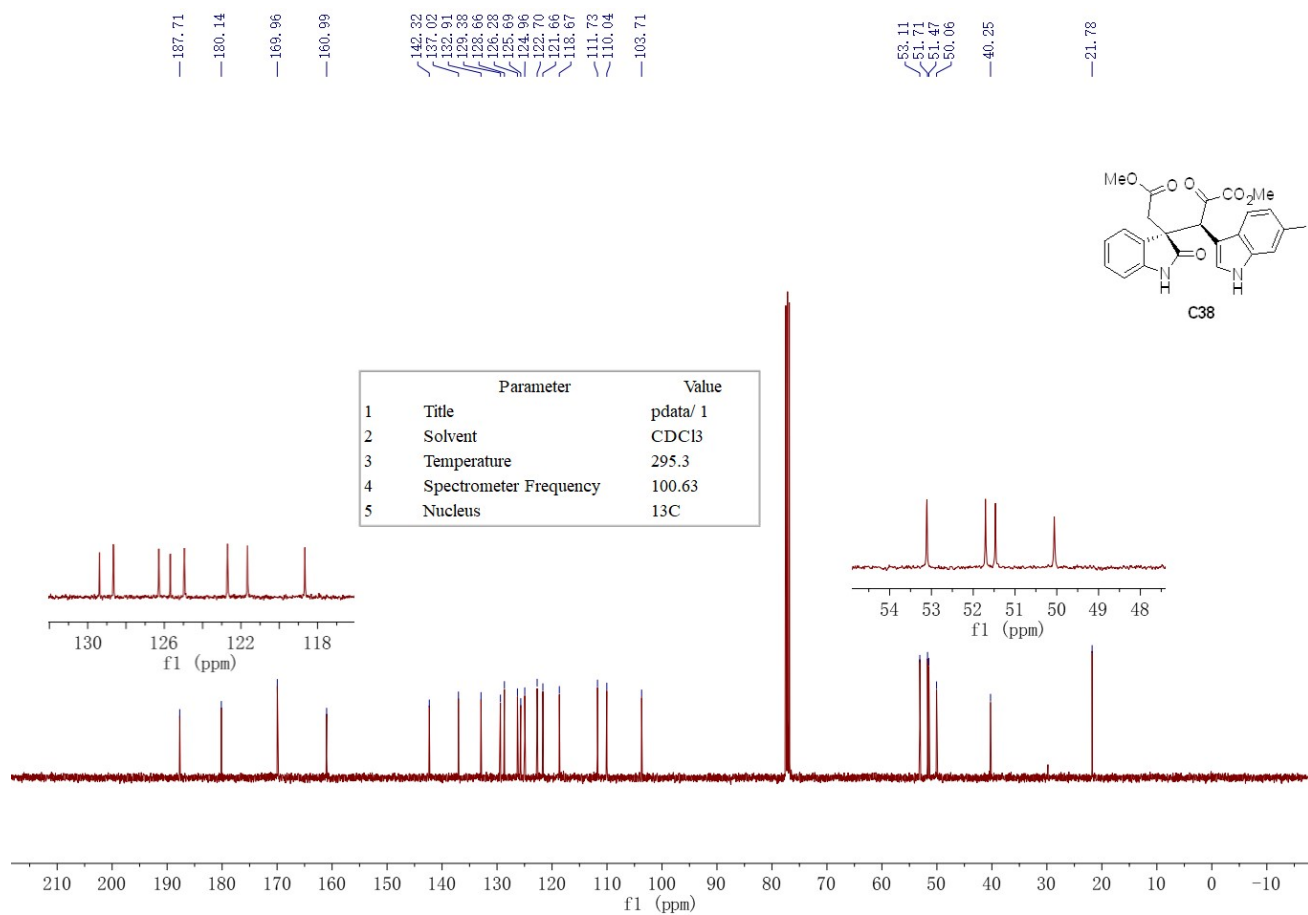
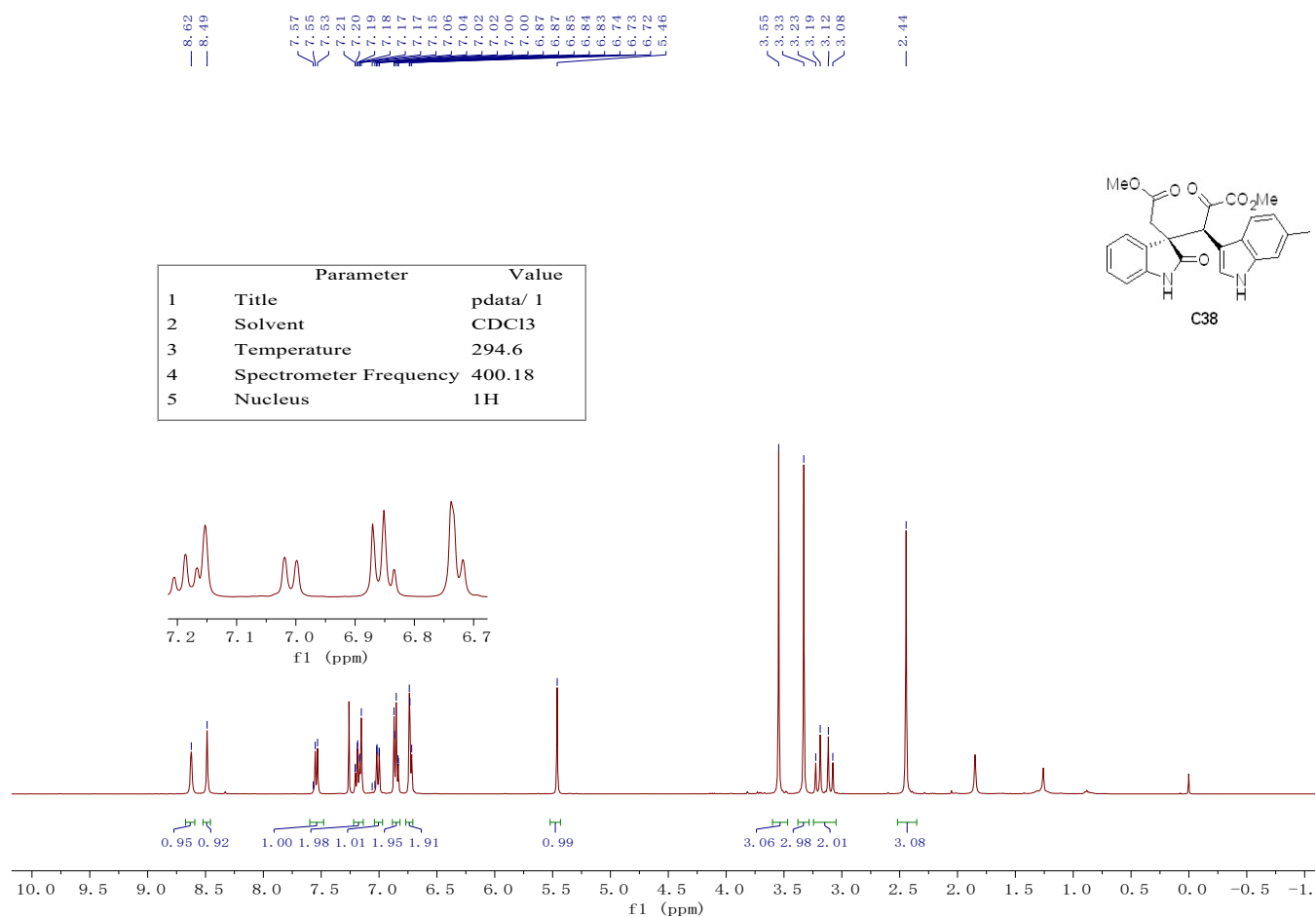


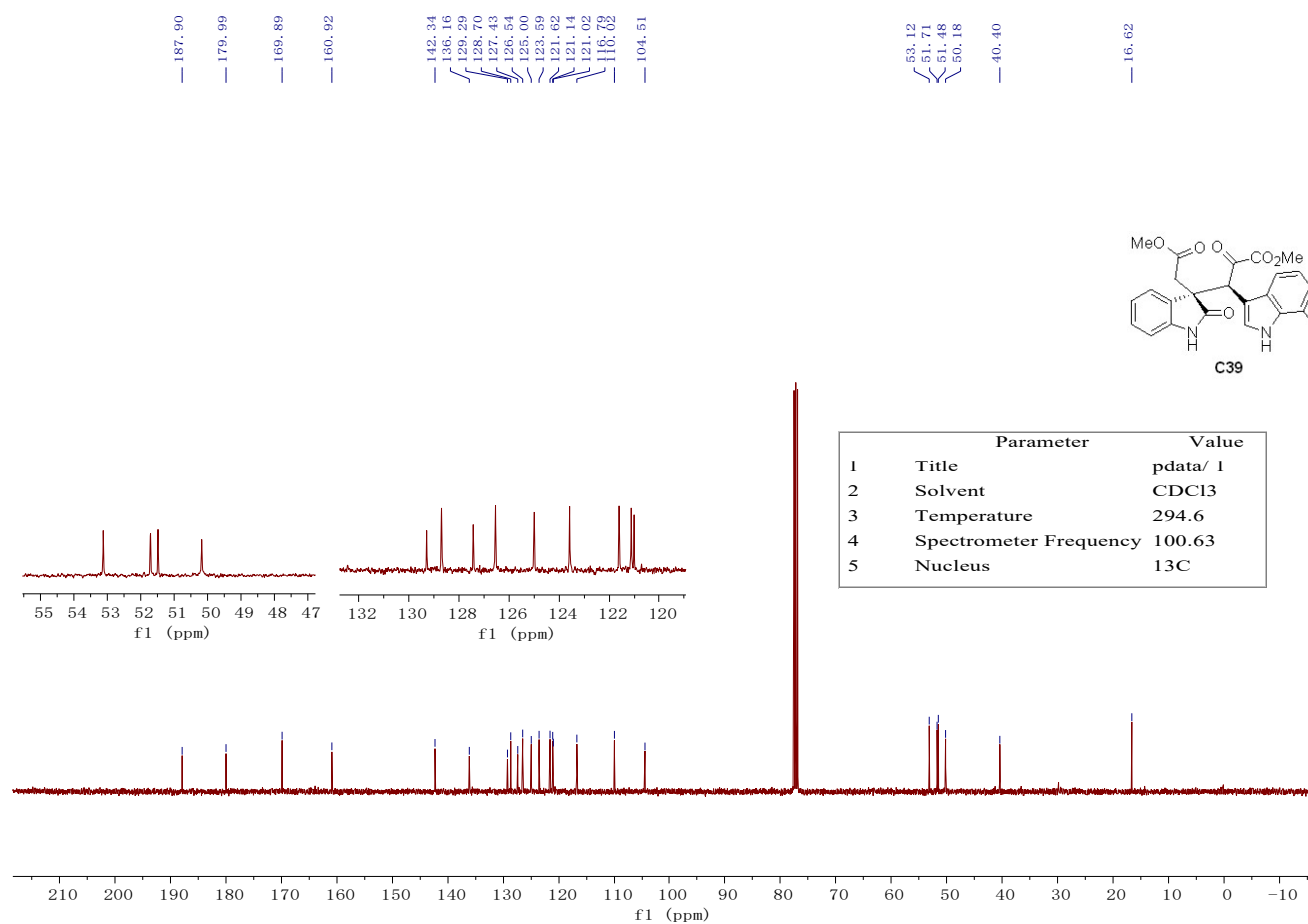
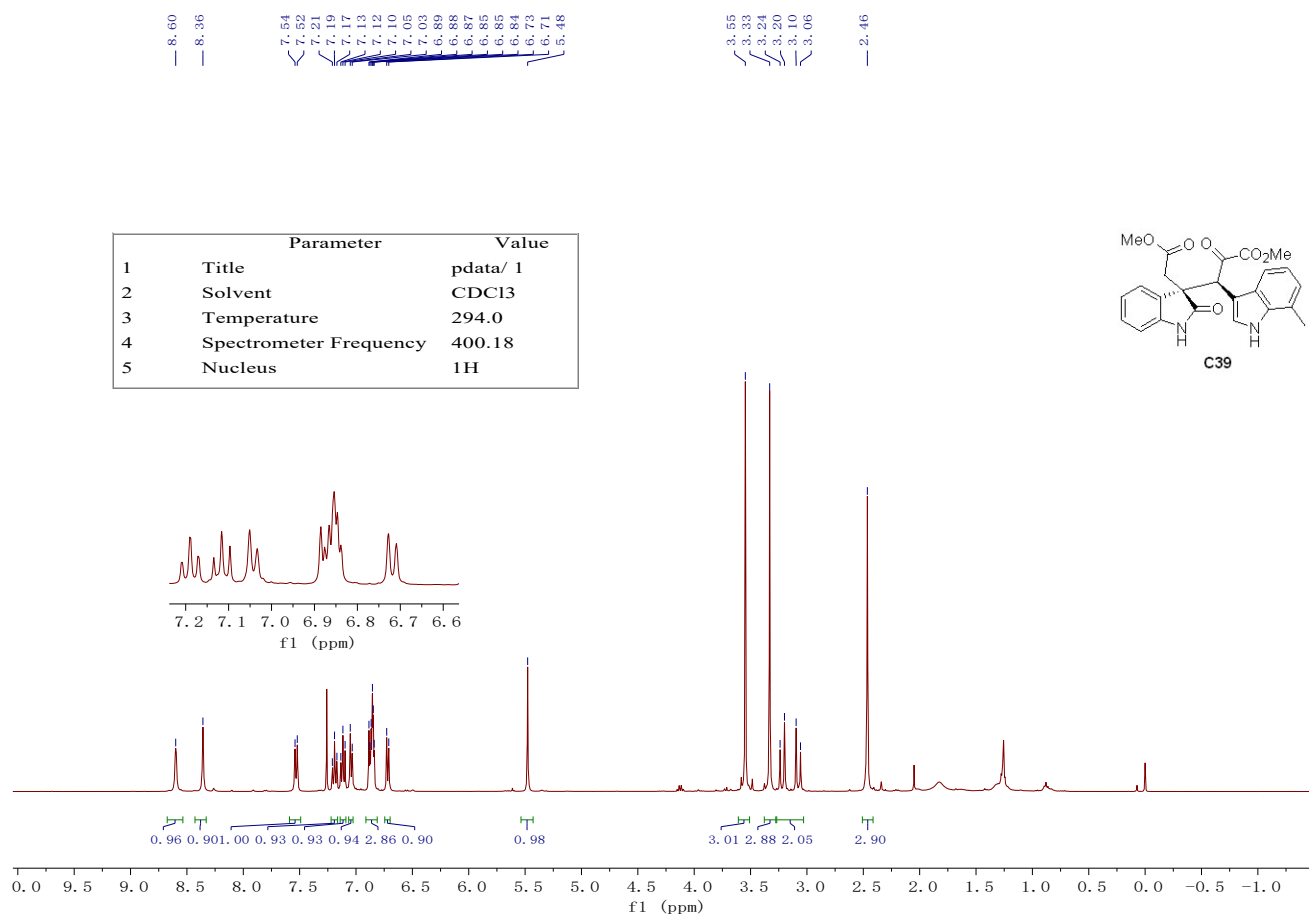
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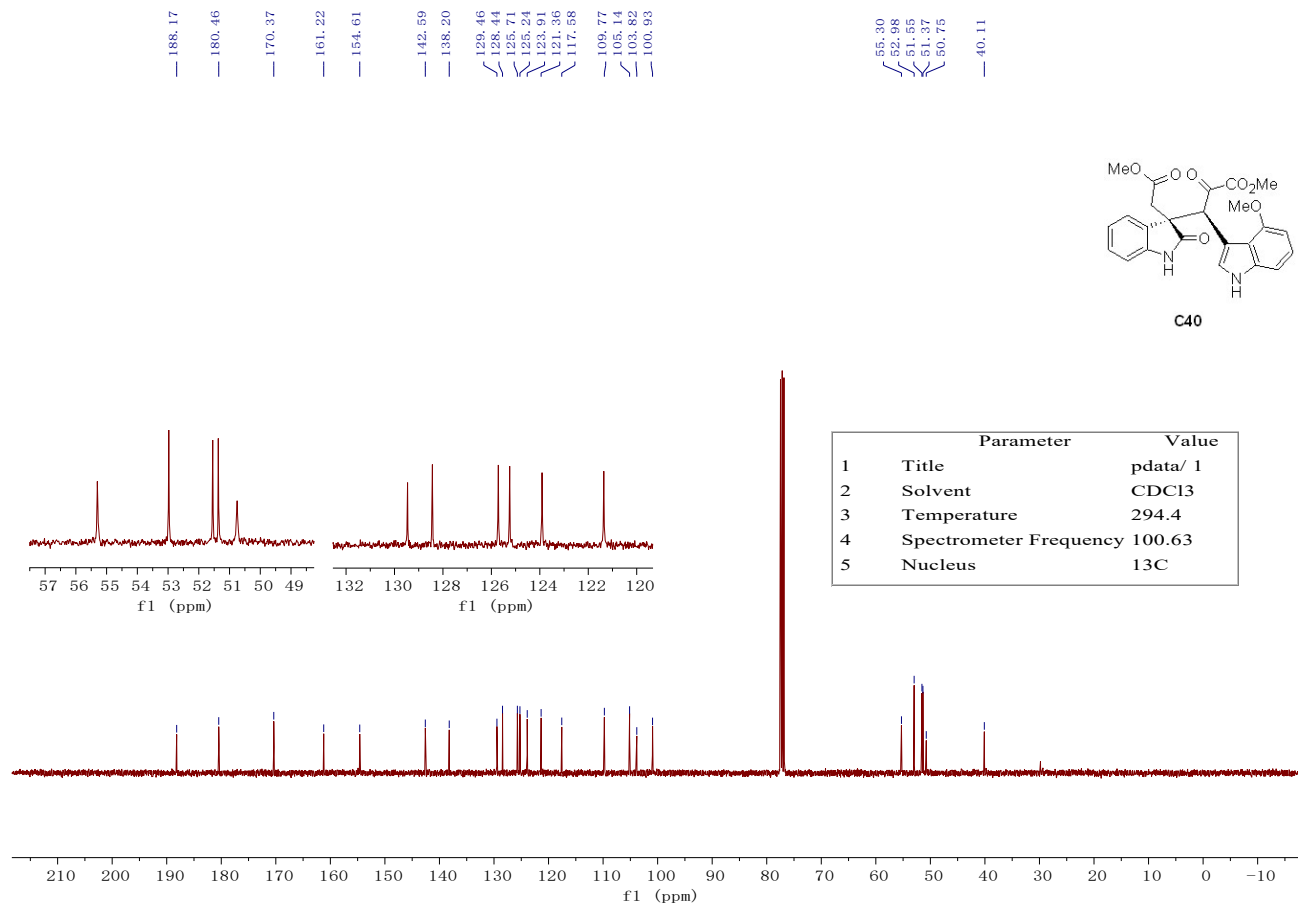
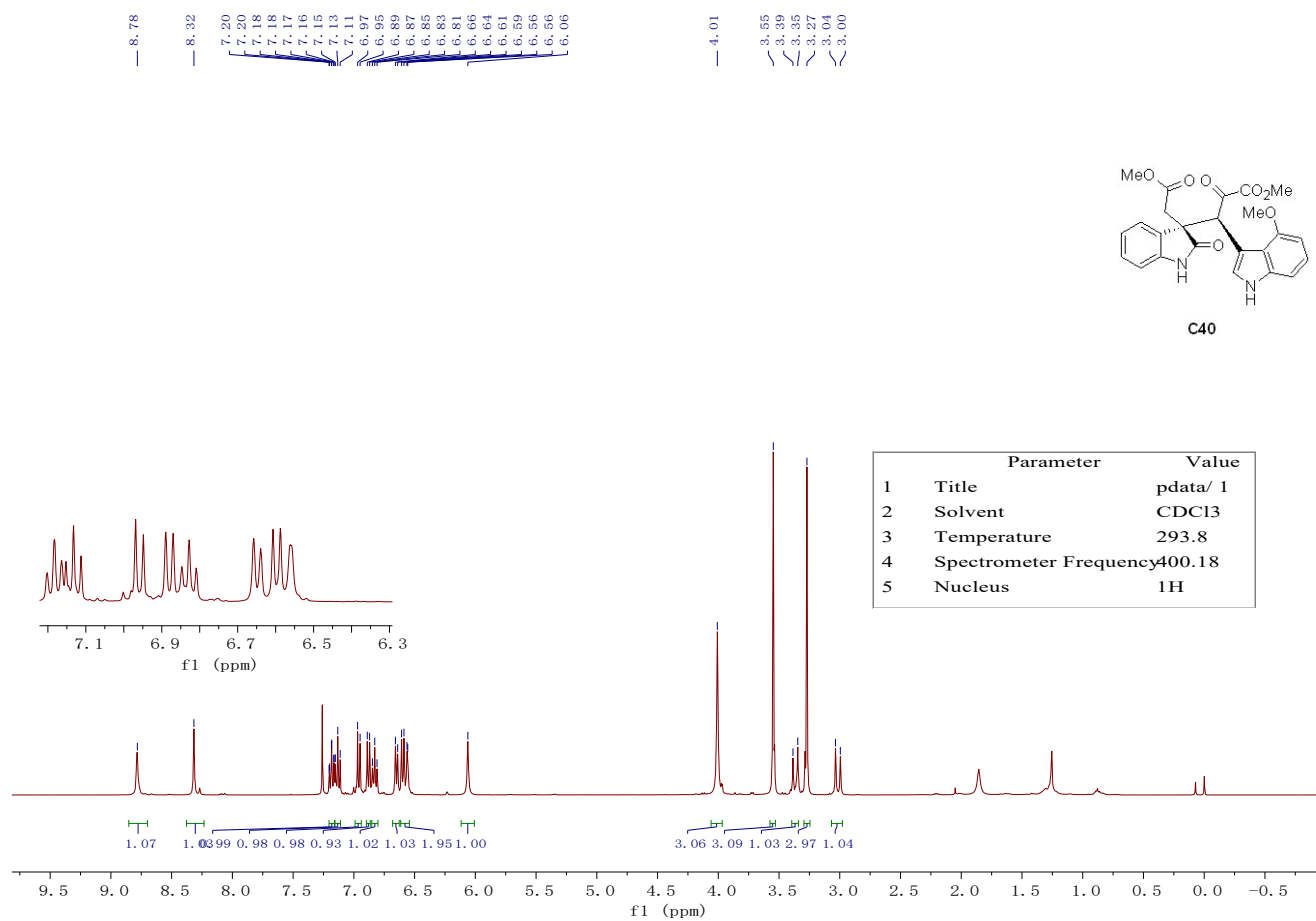


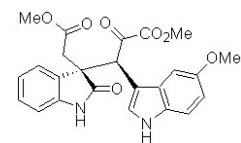
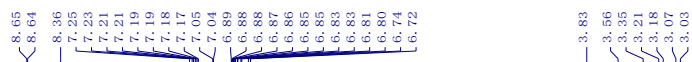




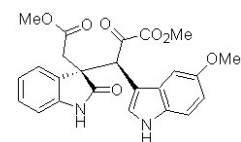
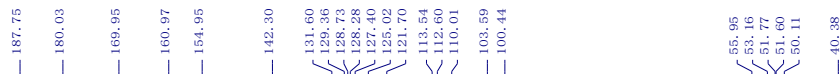
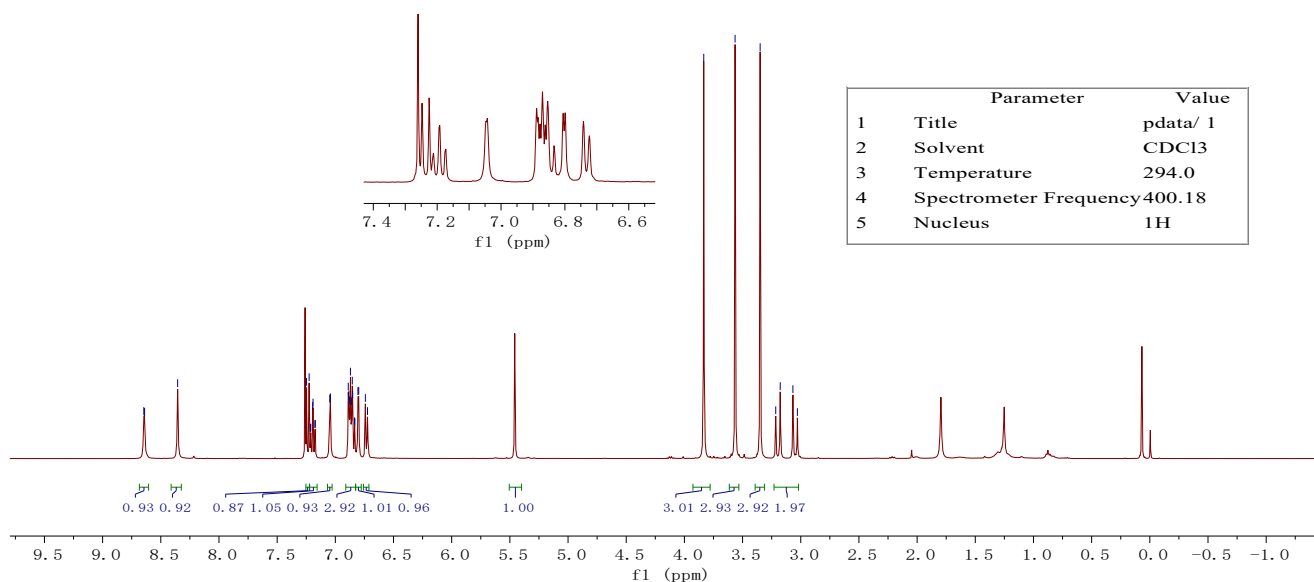




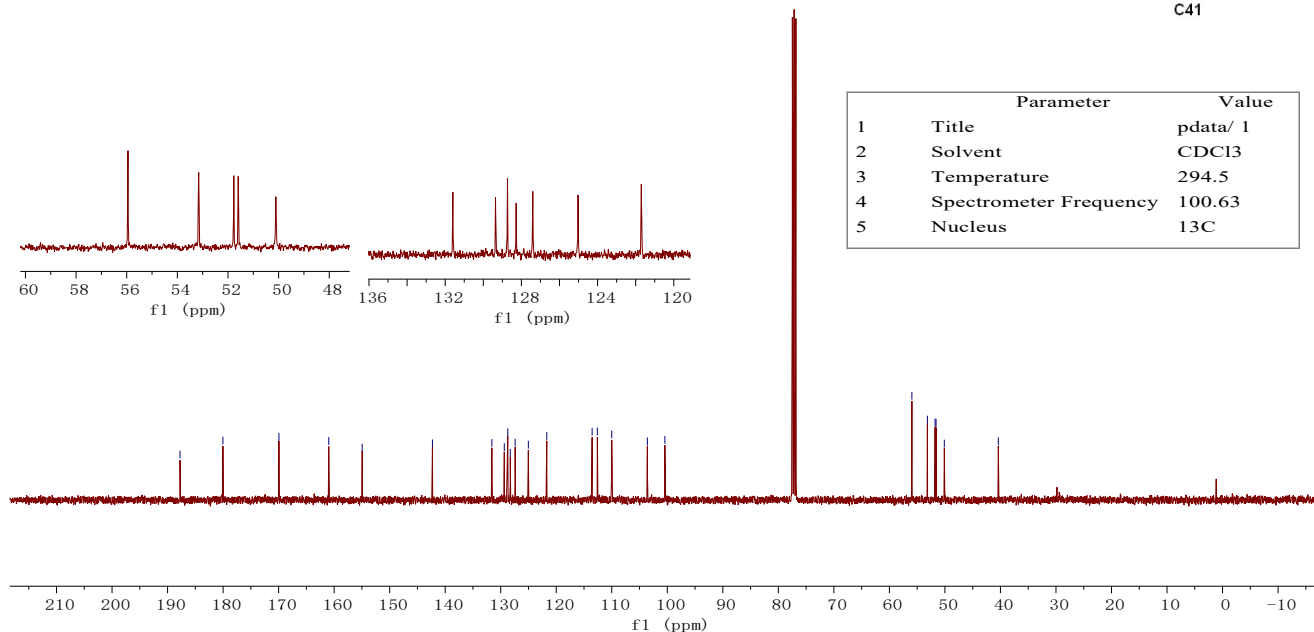


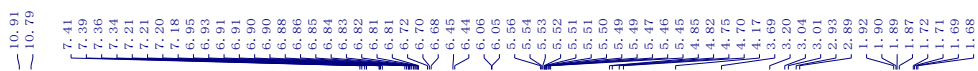


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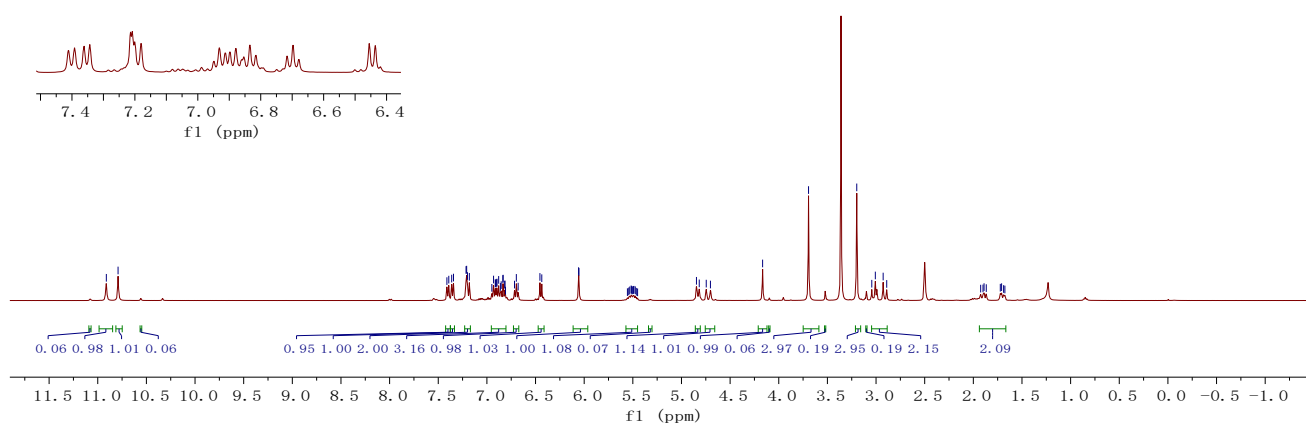
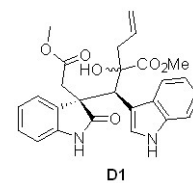


C41

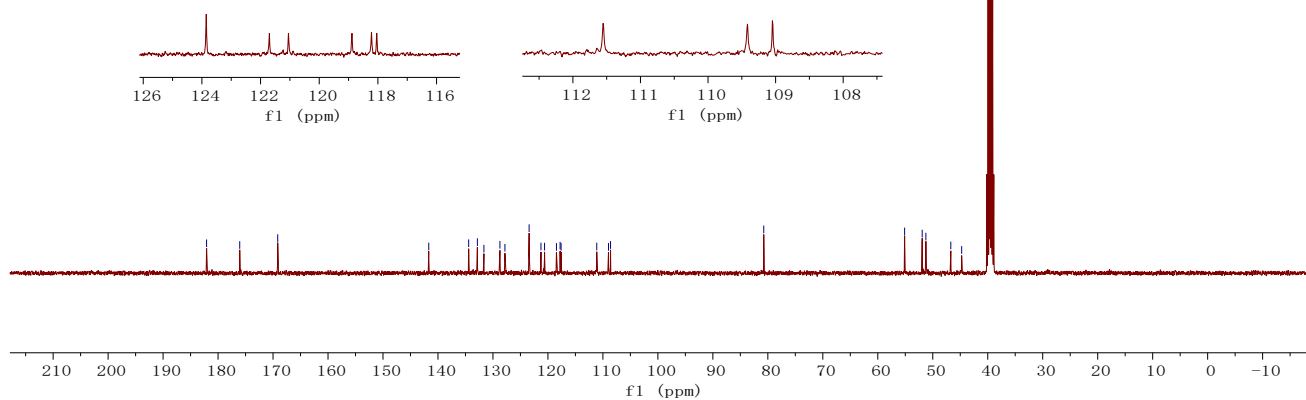
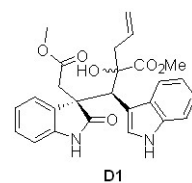




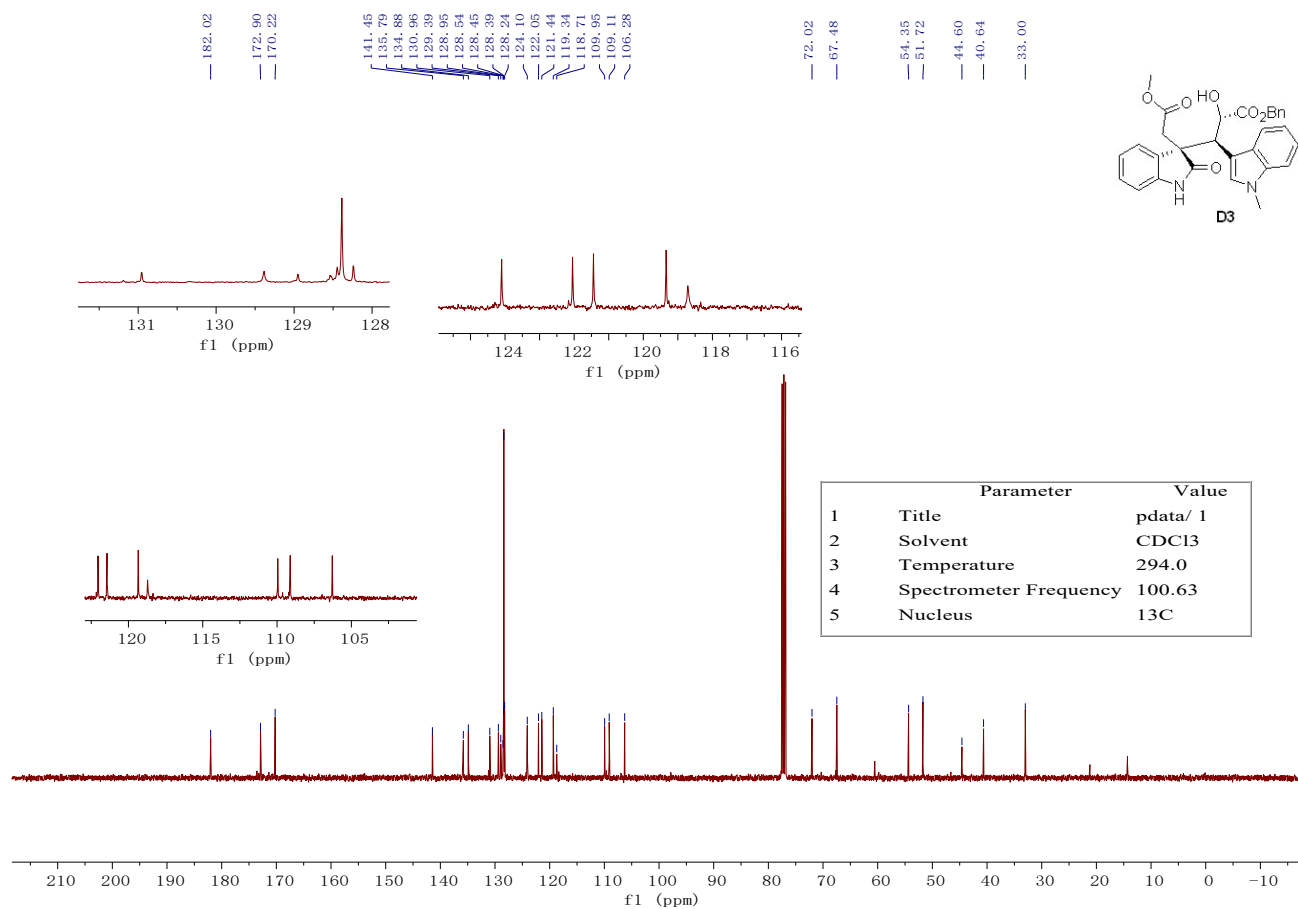
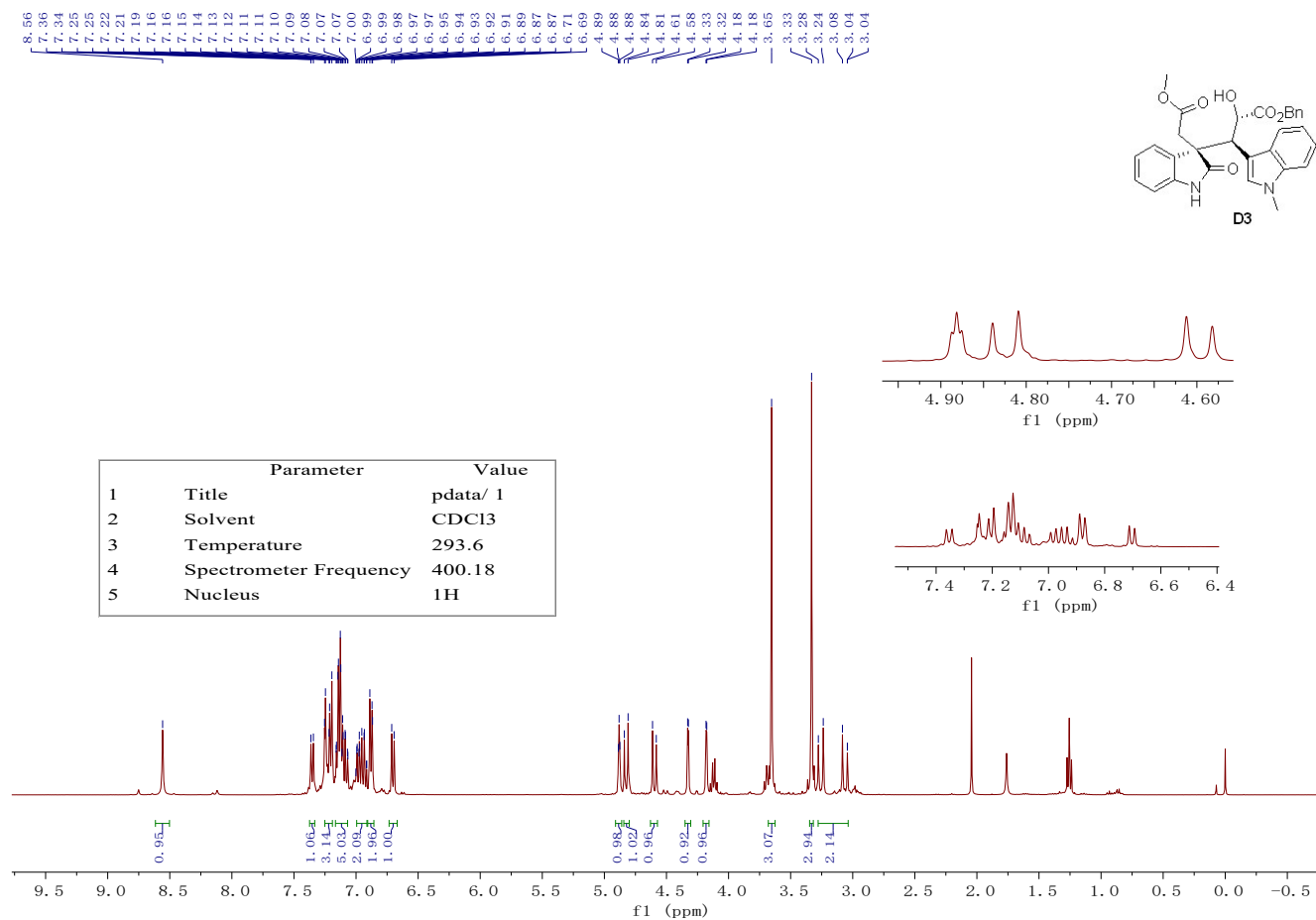
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|---|------------------------|----------|
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| 3 | Temperature            | 293.3    |
| 4 | Spectrometer Frequency | 400.18   |
| 5 | Nucleus                | 1H       |

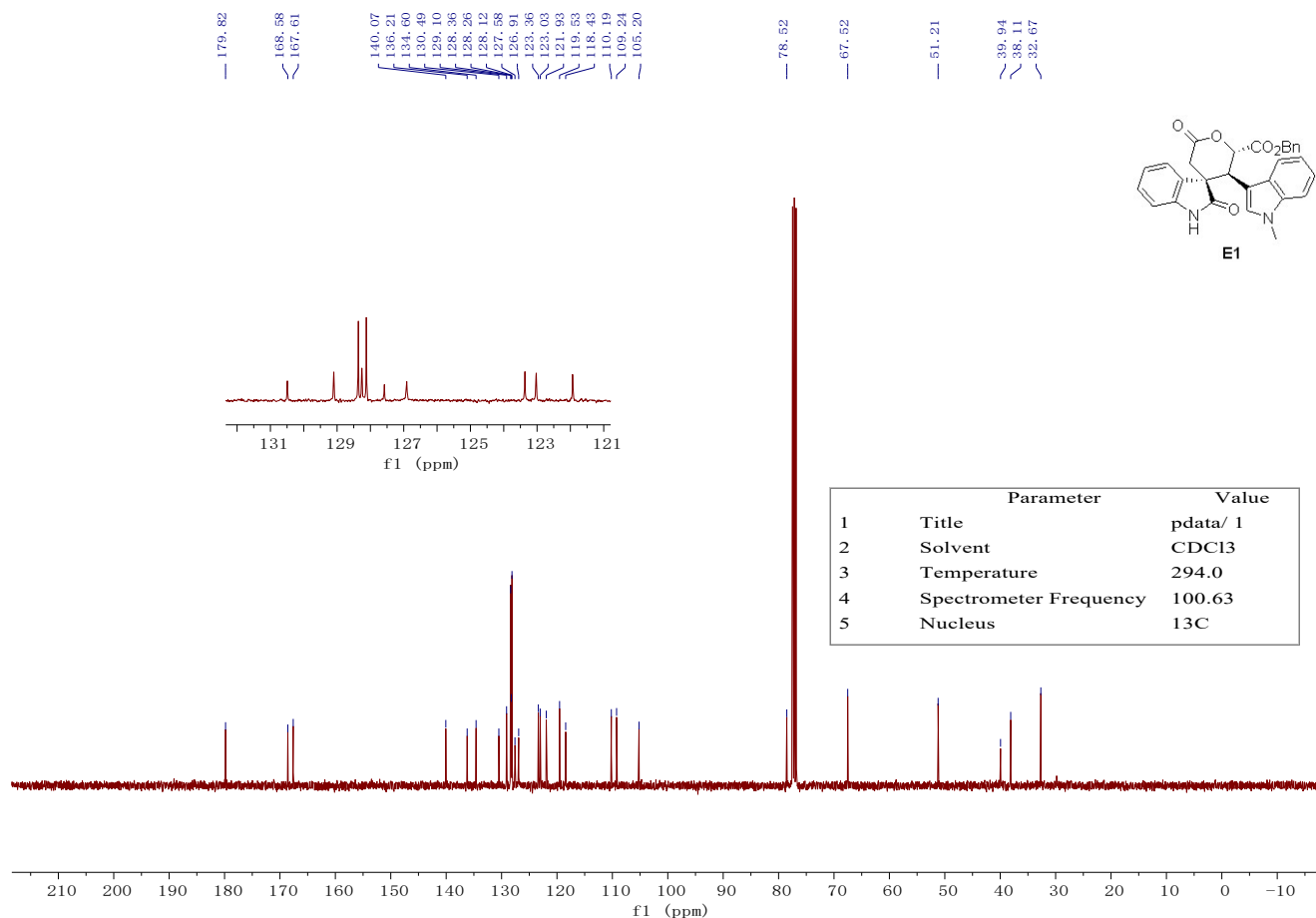
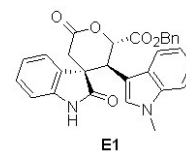
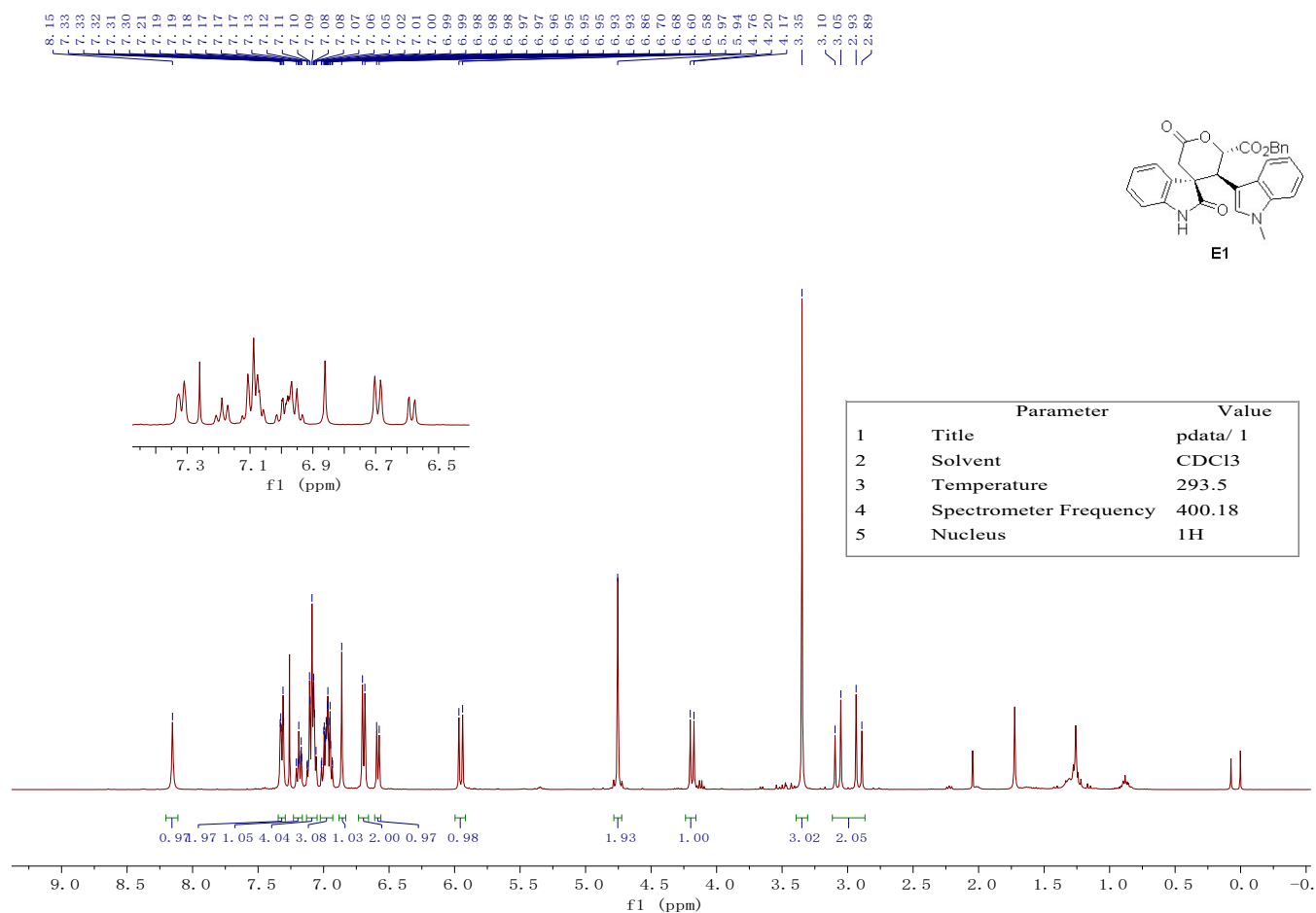
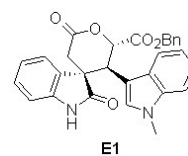


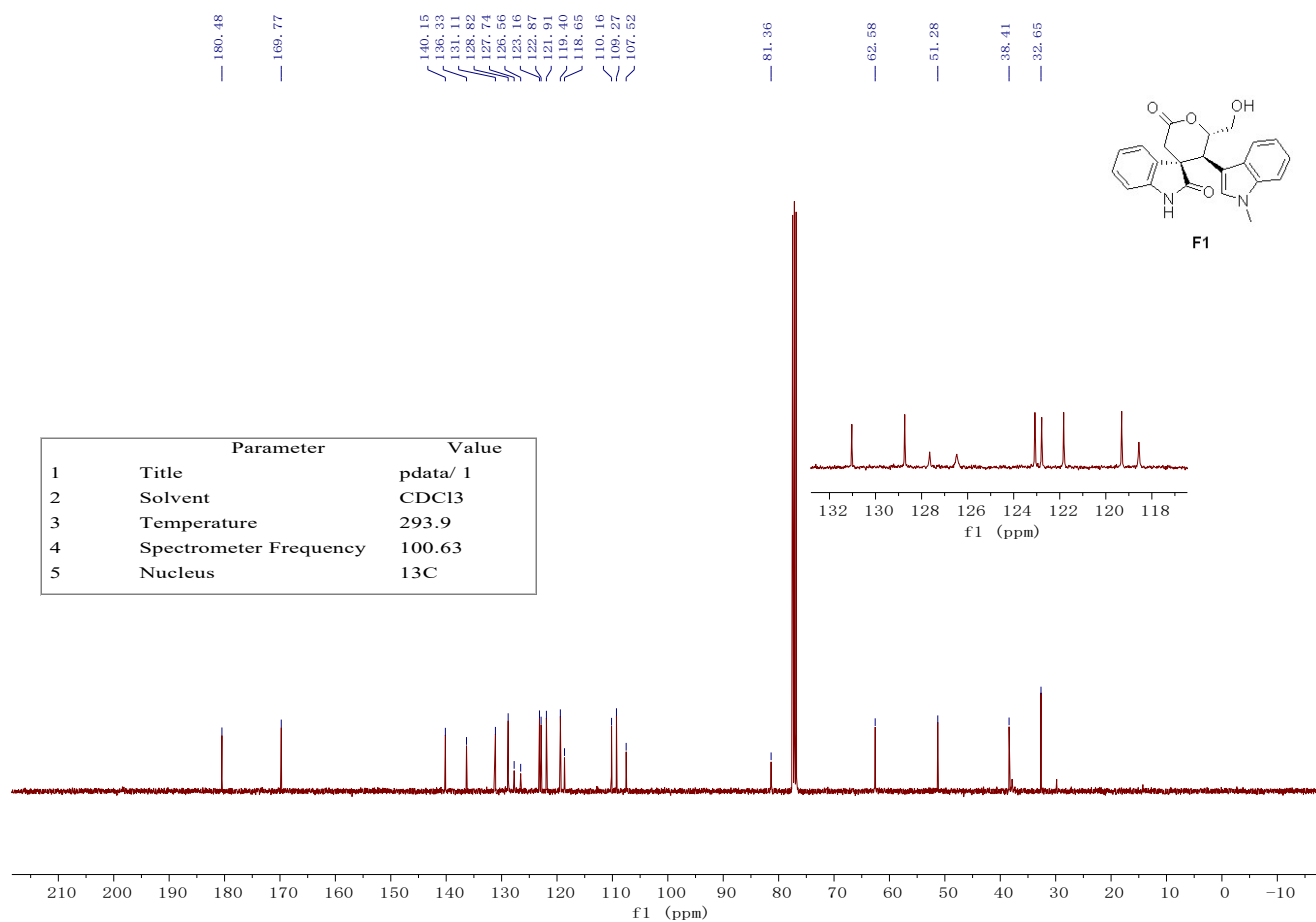
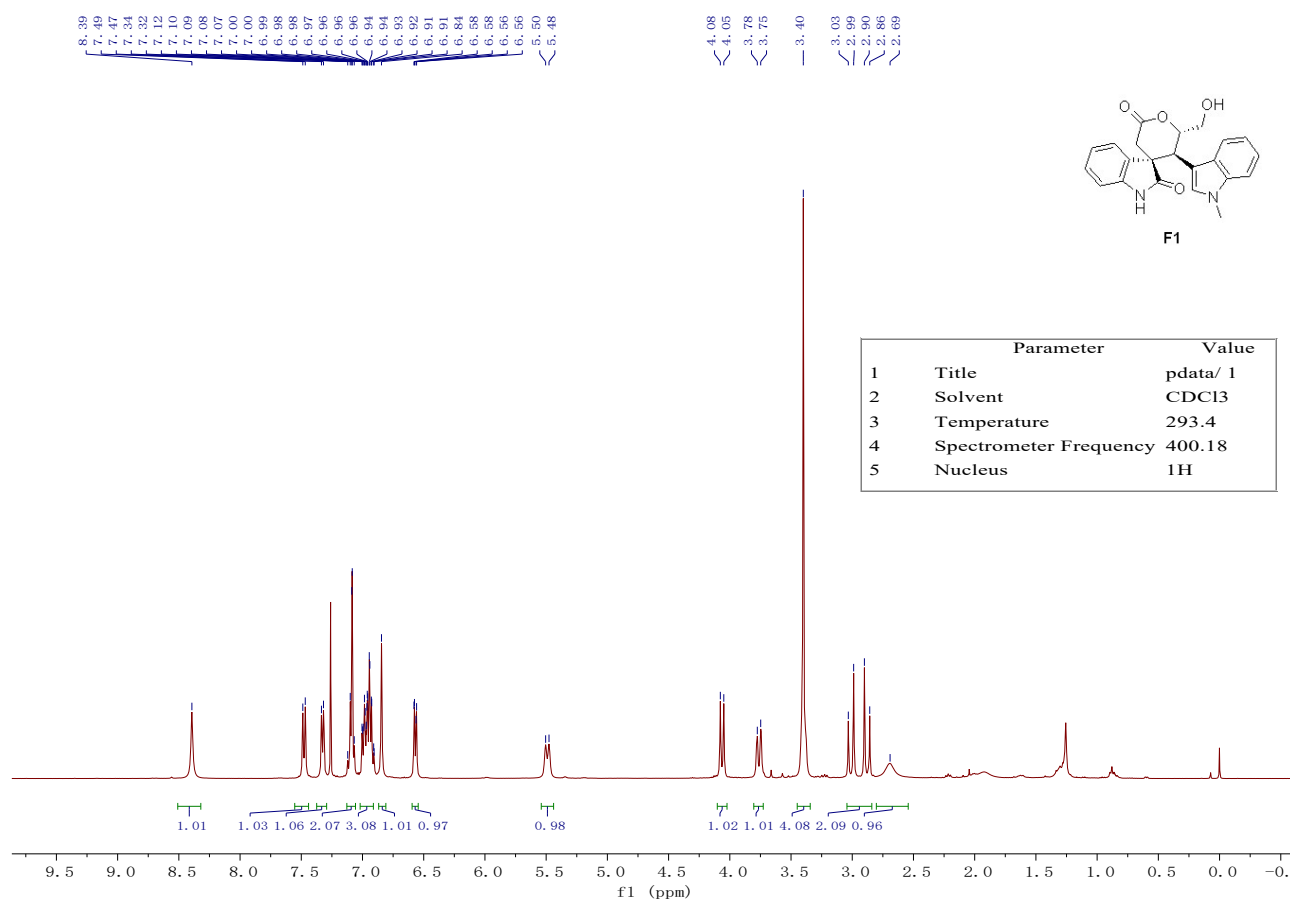
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|---|------------------------|----------|
| 1 | Title                  | pdata/ 1 |
| 2 | Solvent                | DMSO     |
| 3 | Temperature            | 293.7    |
| 4 | Spectrometer Frequency | 100.63   |
| 5 | Nucleus                | 13C      |



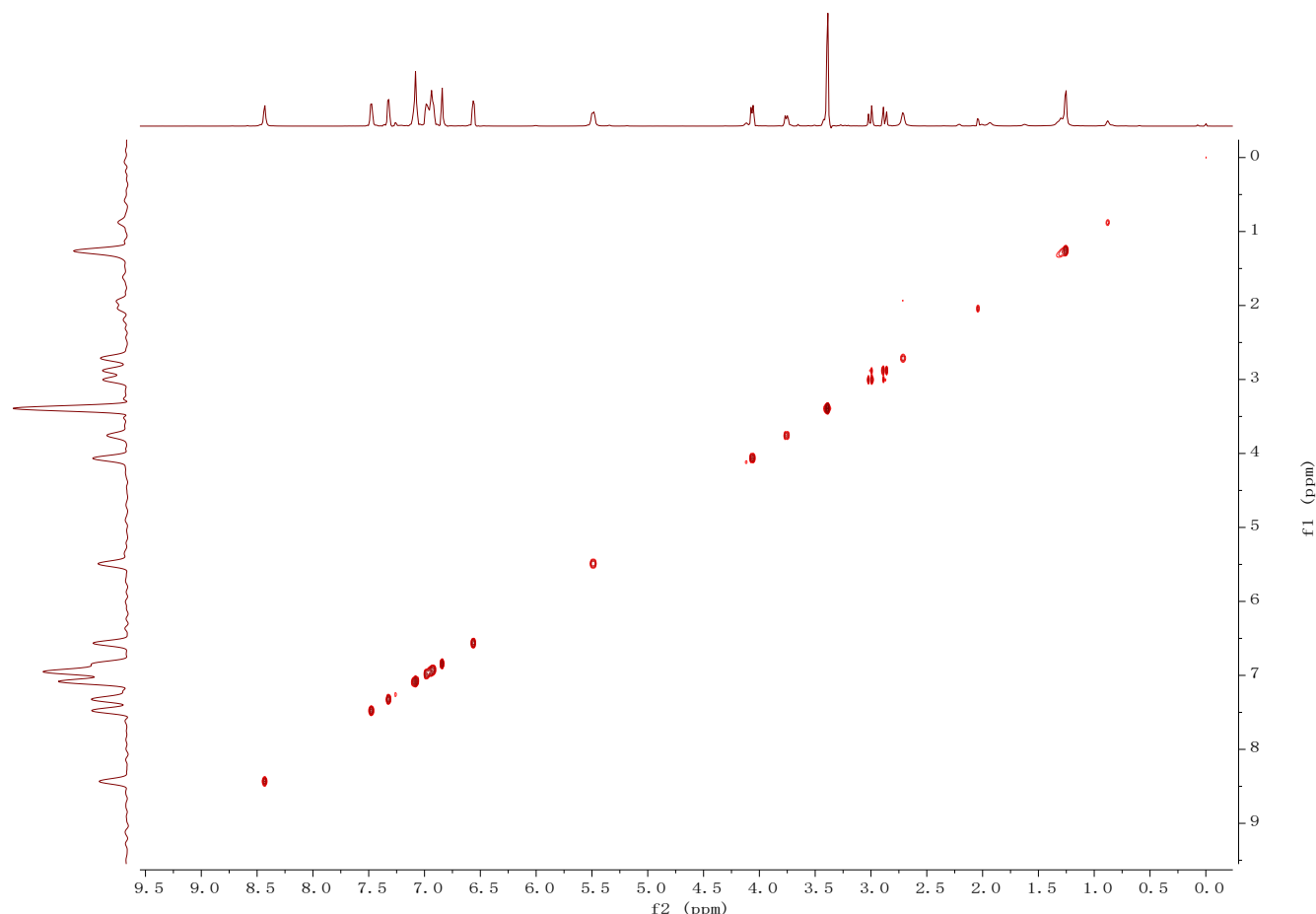








## NOE spectra for F1



## 12 Reference

- [1]. (a) W. K. Yang, P. Dong, J. Xu, J. Yang, X. H. Liu and X. M. Feng, Enantioselective Synthesis of 3-Substituted 3-Amino-2-Oxindoles by Amination with Anilines, *Chem. Eur. J.*, 2021, **27**, 9272–9275; (b) J. Xu, R. Z. Li, N. Xu, X. H. Liu, F. Wang and X. M. Feng, Enantioselective [4 + 2] Cycloaddition/Cyclization Cascade Reaction and Total Synthesis of Cis-Bis(Cyclotryptamine) Alkaloids, *Org. Lett.*, 2021, **23**, 1856–1861.
- [2]. (a) Y. X. Wang, X. Huang, J. L. Huang, Y. Xiong, B. Qin and X. M. Feng, Asymmetric Cyanosilylation of Aldehydes Catalyzed by Novel Organocatalysts, *Synlett*, 2005, **16**, 2445–2448; (b) X. H. Liu, L. L. Lin and X. M. Feng, Chiral *N,N'*-Dioxides: New Ligands and Organocatalysts for Catalytic Asymmetric Reactions, *Acc. Chem. Res.*, 2011, **44**, 574–587; (c) X. H. Liu, L. L. Lin and X. M. Feng, Chiral *N,N'*-dioxide ligands: synthesis, coordination chemistry and asymmetric catalysis, *Org. Chem. Front.*, 2014, **1**, 298–302.
- [3] T. Nishizawa, S. Grüşchow, D. H. E. Jayamaha, C. Nishizawa-Harada and D. H. Sherman, Enzymatic Assembly of the Bis-Indole Core of Rebeccamycin, *J. Am. Chem. Soc.*, 2006, **128**, 724–725.
- [4] (a) G. M. Sheldrick, A Short History of SHELX Acta Cryst, *Acta Cryst.*, 2008, **A64**, 112–122; (b) G. M. Sheldrick, SHELXT – Integrated Space-Group and Crystalstructure Determination Acta Cryst, *Acta Cryst.*, 2015, **A71**, 3–8; (c) G. M. Sheldrick, Crystal Structure Refinement with SHELXL, *Acta Cryst.*, 2015, **C71**, 3–8; (d) O. V. Dolomanov, L. J. Bourhis, R. J. Gildea, J. A. K. Howard and H. Puschmann, OLEX2: A Complete Structure Solution, Refinement and Analysis Program, *J. Appl. Cryst.*, 2009, **42**, 339–341.
- [5] A. L. Spek, Single-Crystal Structure Validation with The Program PLATON, *J. Appl. Cryst.*, 2003, **36**, 7–13.