

Electronic Supplementary Information

Copper(II)-Catalyzed Cross Dehydrogenative Coupling Reaction of *N*-Hydroxyphthalimide with Alkanes and Ethers Via Unactivated C(sp³)-H Activation at Room Temperature

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Contents

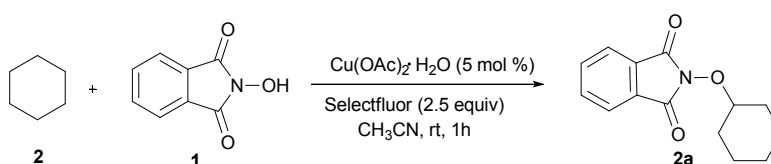
General Experimental Details.....	S2
General Information.....	S2
Typical procedure for the synthesis of 2-(cyclohexyloxy)isoindoline-1,3-dione 2a	S2
Experimental characterization data for compounds.....	S2-S6
NMR spectra of products.....	S7-S26
Electrospray ionization (ESI) spectra of 1-(cyclohexyloxy)-2,2,6,6-tetramethylpiperidine.....	S27

General Experimental Details

General Information:

Unless otherwise stated, all reagents were purchased from commercial suppliers and used without purifications. Melting points are uncorrected. ^1H (400 MHz) NMR and ^{13}C (101 MHz) NMR spectra were recorded on a Varian spectrometer in CDCl_3 using tetramethylsilane (TMS) as internal standards. Chemical shifts (δ) are expressed in ppm and coupling constants J are given in Hz. Mass spectra were measured with a HRMS-APCI instrument or a low-resolution MS instrument using ESI or EI ionization.

Typical experimental procedure for the synthesis of 2-(cyclohexyloxy)isoindoline-1,3-dione **2a**



A flame-dried flask was charged with NHPI (1 mmol, 1 equiv, 0.163 g), cyclohexane (2 mL), F-TEDA- BF_4 (2.5 mmol, 2.5 equiv, 0.885 g), $\text{Cu}(\text{OAc})_2 \cdot \text{H}_2\text{O}$ (5 mol %, 0.01 g), CH_3CN (10 mL) were added and the resulting mixture being stirred at room temperature for 1h. Then aqueous NaHCO_3 solution (10 mL) was added to the reaction mixture, the mixture was extracted with dichloromethane (3 x 10 mL), and the combined organic extracts were successively washed with water (20 mL). The mixture was dried over Na_2SO_4 , and the solvent was removed under vacuum. The crude product was purified by column chromatography on silica gel (petroleum ether/ ethyl acetate 20/1) to afford the corresponding product .

Experimental characterization data for compounds

2-(cyclohexyloxy)isoindoline-1,3-dione (Table 2, **2a**).

60%, 147 mg, white solid, mp 117-118 $^\circ\text{C}$, ^1H NMR (CDCl_3 , 400 MHz): δ 7.83-7.79 (m, 2H), 7.75-7.71 (m, 2H), 4.26-4.19 (m, 1H), 2.04-2.01 (m, 2H), 1.88-1.83 (m, 2H), 1.64-1.52 (m, 3H), 1.34-1.26 (m, 3H).

^{13}C NMR (101MHz, CDCl_3): δ 163.6, 133.8, 128.6, 123.0, 85.6, 31.1, 25.7, 24.1.

HRMS (ESI) m/z : calcd for $\text{C}_{14}\text{H}_{15}\text{NNaO}_3^+$: $M + \text{Na} = 268.0944$; found: 268.0941.

2-(cyclooctyloxy)isoindoline-1,3-dione (Table 2, **2b**).

58%, 158 mg, white solid, mp 105-108 $^\circ\text{C}$, ^1H NMR (CDCl_3 , 400 MHz): δ 7.82-7.79 (m, 2H), 7.75-7.71 (m, 2H), 4.43-4.37 (m, 1H), 2.04-1.88 (m, 4H), 1.85-1.77 (m, 2H), 1.63-1.43 (m, 8H).

^{13}C NMR (101 MHz, CDCl_3): δ 164.2, 134.2, 128.9, 123.2, 88.7, 30.1, 27.2, 25.4, 23.0.

HRMS (ESI) m/z : calcd for $\text{C}_{16}\text{H}_{19}\text{NNaO}_3^+$: $M + \text{Na} = 296.1257$; found: 296.1266.

2-(cyclopentyloxy)isoindoline-1,3-dione (Table 2, **2c**).

42%, 97 mg, white solid, mp 81-85 °C, ¹H NMR (CDCl₃, 400 MHz): δ 7.83-7.81 (m, 2H), 7.75-7.72 (m, 2H), 4.93-4.91 (m, 1H), 2.00-1.90 (m, 4H), 1.81-1.72 (m, 2H), 1.68-1.58 (m, 2H).

¹³C NMR (101 MHz, CDCl₃): δ 164.0, 134.2, 128.9, 123.2, 90.3, 31.6, 23.6.

HRMS (ESI) m/z: calcd for C₁₃H₁₃NNaO₃⁺: M + Na = 254.0788; found: 254.0799.

2-(cyclododecyloxy)isoindoline-1,3-dione (Table 2, 2d).

50%, 165 mg, white solid, mp 111-112 °C, ¹H NMR (CDCl₃, 400 MHz): δ 7.81-7.75 (m, 2H), 7.70-7.66 (m, 2H), 4.40-4.34 (m, 1H), 1.87-1.79 (m, 2H), 1.70-1.59 (m, 4H), 1.46-1.25 (m, 16H).

¹³C NMR (101 MHz, CDCl₃): δ 163.6, 133.8, 128.6, 122.9, 86.3, 28.9, 24.7, 24.1, 23.9, 23.8, 21.6.

HRMS (ESI) m/z: calcd for C₂₀H₂₇NNaO₃⁺: M + Na = 352.1883; found: 352.1889.

2-((1r,3r,5r,7r)-adamantan-2-yloxy)isoindoline-1,3-dione (Table 2, 2e).

37%, 110 mg, white solid, mp 175-177 °C, ¹H NMR (CDCl₃, 400 MHz): δ 7.81-7.78 (m, 2H), 7.74-7.70 (m, 2H), 4.33 (s, 1H), 2.44 (d, *J* = 12.8 Hz, 2H), 2.23 (d, *J* = 1.6 Hz, 2H), 1.91-1.84 (m, 4H), 1.75 (s, 2H), 1.67 (d, *J* = 12.8 Hz, 2H), 1.61-1.57 (m, 2H).

¹³C NMR (101 MHz, CDCl₃): δ 163.5, 133.8, 128.6, 122.9, 90.2, 37.6, 36.9, 31.9, 31.7, 27.7, 27.6.

HRMS (ESI) m/z: calcd for C₁₈H₁₉NNaO₃⁺: M + Na = 320.1257; found: 320.1249.

2-((tetrahydrofuran-2-yl)oxy)isoindoline-1,3-dione (Table 2, 2f).

89%, 207 mg, white solid, mp 133-134 °C, ¹H NMR (CDCl₃, 400 MHz): δ 7.82-7.78 (m, 2H), 7.73-7.69 (m, 2H), 5.77 (d, *J* = 4.8 Hz, 1H), 4.37-4.31 (m, 1H), 4.03-4.00 (m, 1H), 2.34-2.20 (m, 2H), 2.15-2.10 (m, 1H), 2.00-1.90 (m, 1H).

¹³C NMR (101 MHz, CDCl₃): δ 163.6, 134.1, 128.9, 123.2, 108.7, 69.1, 30.9, 22.6.

HRMS (ESI) m/z: calcd for C₁₂H₁₁NNaO₄⁺: M + Na = 256.0580; found: 256.0581.

2-((5-methyltetrahydrofuran-2-yl)oxy)isoindoline-1,3-dione (Table 2, 2g).

70%, 173 mg, white solid, mp 125-126 °C, ¹H NMR (CDCl₃, 400 MHz): δ 7.81-7.79 (m, 2H), 7.72-7.70 (m, 2H), 5.75 (d, *J* = 12.4 Hz, 1H), 4.74-4.66 (m, 1H), 2.29-2.22 (m, 3H), 1.55-1.47 (m, 1H), 1.25 (d, *J* = 6.4 Hz, 3H).

¹³C NMR (101 MHz, CDCl₃): δ 163.4, 134.0, 129.0, 123.2, 108.8, 76.5, 30.7, 30.3, 20.6.

HRMS (ESI) m/z: calcd for C₁₃H₁₃NNaO₄⁺: M + Na = 270.0737; found: 270.0736.

2-((tetrahydro-2H-pyran-2-yl)oxy)isoindoline-1,3-dione (Table 2, 2h).

71%, 175 mg, white solid, mp 122-124 °C, ¹H NMR (CDCl₃, 400 MHz): δ 7.81-7.80 (m, 2H), 7.72-7.71 (m, 2H), 5.41 (s, 1H), 4.54-4.48 (m, 1H), 3.65 (d, *J* = 10.4 Hz, 1H), 2.11 (d, *J* = 13.6 Hz, 1H), 2.01-1.91 (m, 1H), 1.87-1.69 (m, 4H).

¹³C NMR (101 MHz, CDCl₃): δ 163.5, 134.1, 129.0, 123.2, 102.9, 62.24, 27.8, 24.9, 17.7.

HRMS (ESI) m/z: calcd for C₁₃H₁₃NNaO₄⁺: M + Na = 270.0737; found: 270.0749.

2-((1,4-dioxan-2-yl)oxy)isoindoline-1,3-dione (Table 2, 2i).

78%, 194 mg, white solid, mp 186-188 °C, ¹H NMR (CDCl₃, 400 MHz): δ 7.84-7.80 (m, 2H), 7.77-7.73 (m, 2H), 5.23 (d, *J* = 1.2 Hz, 1H), 4.87-4.41 (m, 1H), 4.15 (d, *J* = 10.4 Hz, 1H), 3.91-3.87(m, 1H), 3.83-3.79 (m, 2H), 3.54 (dd, *J*₁ = 2Hz, *J*₂ = 11.6 Hz, 1H).

¹³C NMR (101 MHz, CDCl₃): δ 163.2, 134.3, 128.9, 123.4, 99.4, 66.1, 65.9, 60.9.

HRMS (ESI) *m/z*: calcd for C₁₂H₁₁NNaO₅⁺: *M* + Na = 272.0529; found: 272.0538.

2-(1-ethoxyethoxy)isoindoline-1,3-dione (Table 2, 2j).

58%, 136 mg, white oil, ¹H NMR (CDCl₃, 400 MHz): δ 7.78-7.74 (m, 2H), 7.72-7.68 (m, 2H), 5.26 (q, *J* = 5.2 Hz, 1H), 4.18-4.11 (m, 1H), 3.84-3.76 (m, 1H), 1.52 (d, *J* = 5.2 Hz, 3H), 1.23 (t, *J* = 7.2Hz, 3H).

¹³C NMR (101 MHz, CDCl₃): δ 163.2, 133.7, 128.5, 122.7, 106.0, 63.5, 19.3, 15.3.

HRMS (ESI) *m/z*: calcd for C₁₂H₁₃NNaO₄⁺: *M* + Na = 258.0737; found: 258.0749.

2-(2-chloro-1-(2-chloroethoxy)ethoxy)isoindoline-1,3-dione (Table 2, 2k).

70%, 212 mg, white solid, mp 84-85 °C, ¹H NMR (CDCl₃, 400 MHz): δ 7.85-7.83 (m, 2H), 7.79-7.76 (m, 2H), 5.23 (dd, *J*₁ = 3.2 Hz, *J*₂ = 8 Hz, 1H), 4.49-4.32 (m, 1H), 4.16-4.10 (m, 1H), 3.92(dd, *J*₁ = 2.8 Hz, *J*₂ = 11.6 Hz, 1H), 3.72 (t, *J* = 6.0 Hz, 2H), 3.66 (dd, *J*₁ = 7.6 Hz, *J*₂ = 11.6 Hz, 1H).

¹³C NMR (101 MHz, CDCl₃): δ 163.2, 134.1, 128.4, 123.3, 107.8, 70.1, 42.8, 42.6.

HRMS (ESI) *m/z*: calcd for C₁₂H₁₁Cl₂NNaO₄⁺: *M* + Na = 325.9957; found: 325.9950.

2-(2-bromo-1-(2-bromoethoxy)ethoxy)isoindoline-1,3-dione (Table 2, 2l).

76%, 296 mg, white solid, mp 110-112 °C, ¹H NMR (CDCl₃, 400 MHz): δ 7.86-7.82 (m, 2H), 7.80-7.76 (m, 2H), 5.26 (dd, *J*₁ = 2.8 Hz, *J*₂ = 8.0 Hz, 1H), 4.55-4.49 (m, 1H), 4.21-4.15 (m, *J* = 10.4 Hz, 1H), 3.75 (dd, *J*₁ = 2.8 Hz, *J*₂ = 10.8 Hz, 1H), 3.59-3.53 (m, 2H), 3.50 (dd, *J*₁ = 8.4 Hz, *J*₂ = 11.2 Hz, 1H).

¹³C NMR (101 MHz, CDCl₃): δ 163.2, 134.1, 128.4, 123.3, 107.8, 70.0, 29.9, 29.8.

HRMS (ESI) *m/z*: calcd for C₁₂H₁₁Br₂NNaO₄⁺: *M* + Na = 413.8947; found: 413.8927.

2-(1-butoxybutoxy)isoindoline-1,3-dione (Table 2, 2m).

45%, 131 mg, white oil, ¹H NMR (CDCl₃, 400 MHz): δ 7.82-7.78 (m, 2H), 7.77-7.73 (m, 2H), 5.10 (dd, *J*₁ = 4.4 Hz, *J*₂ = 7.2 Hz, 1H), 4.55-4.49 (m, 1H), 4.09-4.03 (m, 1H), 3.78-3.73 (m, 1H), 1.95-1.87 (m, 1H), 1.77-1.71 (m, 1H), 1.62-1.47 (m, 4H), 1.43-1.34 (m, 1H), 0.99-0.90 (m, 6H).

¹³C NMR (101 MHz, CDCl₃): δ 162.9, 133.5, 128.4, 122.5, 109.4, 68.0, 34.8, 31.8, 19.3, 18.1, 14.0.

HRMS (ESI) *m/z*: calcd for C₁₆H₂₁NNaO₄⁺: *M* + Na = 314.1363; found: 314.1372.

2-((cyclopentyloxy)methoxy)isoindoline-1,3-dione (Table 2, 2n).

40%, 104 mg, white solid, mp 72-73 °C, ¹H NMR (CDCl₃, 400 MHz): δ 7.83-7.79 (m, 2H), 7.75-

7.70 (m, 2H), 5.16 (s, 2H), 4.78-4.70 (m, 1H), 1.88-1.81 (m, 2H), 1.73-1.65 (m, 4H), 1.62-1.54 (m, 2H).

¹³C NMR (101 MHz, CDCl₃): δ 162.8, 133.8, 128.6, 123.0, 97.6, 80.2, 32.5, 23.8.

HRMS (ESI) m/z: calcd for C₁₄H₁₅NNaO₄⁺: M + Na = 284.0893, found: 284.0881.

2-(tert-butoxymethoxy)isoindoline-1,3-dione (Table 2, 2o).

61%, 152 mg, white solid, mp 95-96 °C, ¹H NMR (CDCl₃, 400 MHz): δ 7.83-7.79 (m, 2H), 7.74-7.70 (m, 2H), 5.25 (s, 2H), 1.38 (s, 9H).

¹³C NMR (101 MHz, CDCl₃): δ 162.9, 133.7, 128.6, 122.9, 94.8, 77.1, 28.7.

HRMS (ESI) m/z: calcd for C₁₃H₁₅NNaO₄⁺: M + Na = 272.0893; found: 272.0898.

2-(1-(tert-butoxy)ethoxy)isoindoline-1,3-dione (Table 2, 2p).

30%, 79 mg, white solid, mp 68-70 °C, ¹H NMR (CDCl₃, 400 MHz): δ 7.83-7.81 (m, 2H), 7.46-7.34 (m, 2H), 5.54 (d, *J* = 5.2 Hz, 1H), 1.46 (d, *J* = 4.8 Hz, 3H), 1.31 (s, 9H).

¹³C NMR (101 MHz, CDCl₃): δ 164.2, 134.2, 129.0, 123.2, 101.0, 76.3, 28.4, 21.1.

HRMS (ESI) m/z: calcd for C₁₄H₁₇NNaO₄⁺: M + Na = 286.1050; found: 286.1057.

2-((2-methoxyethoxy)methoxy)isoindoline-1,3-dione (Table 2, 2q).

29%, 72 mg, white solid, mp 80-82 °C, ¹H NMR (CDCl₃, 400 MHz): δ 7.78-7.76 (m, 2H), 7.70-7.68 (m, 2H), 5.17 (s, 2H), 4.10 (t, *J* = 4.4 Hz, 2H), 3.59 (t, *J* = 4.4 Hz, 2H), 3.36 (s, 3H).

¹³C NMR (101 MHz, CDCl₃): δ 162.7, 133.8, 128.6, 123.0, 99.8, 71.5, 69.1, 59.1.

HRMS (ESI) m/z: calcd for C₁₂H₁₃NNaO₅⁺: M + Na = 274.0686; found: 274.0697.

2-(1,2-dimethoxyethoxy)isoindoline-1,3-dione (Table 2, 2r).

31%, 78 mg, white solid, mp 65-66 °C, ¹H NMR (CDCl₃, 400 MHz): δ 7.79-7.61 (m, 2H), 7.72-7.69 (m, 2H), 5.10 (dd, *J*₁ = 4.0 Hz, *J*₂ = 6.0 Hz, 1H), 3.77 (dd, *J*₁ = 4.0 Hz, *J*₂ = 10.8 Hz, 1H), 3.74 (s, 3H), 3.58 (dd, *J*₁ = 6.0 Hz, *J*₂ = 10.8 Hz, 1H), 3.39 (s, 3H).

¹³C NMR (101 MHz, CDCl₃): δ 163.1, 133.9, 128.6, 123.1, 108.1, 71.8, 59.6, 57.2.

HRMS (ESI) m/z: calcd for C₁₂H₁₃NNaO₅⁺: M + Na = 274.0686; found: 274.0684.

2-((tetrahydrothiophen-2-yl)oxy)isoindoline-1,3-dione (Table 2, 2s).

88%, 219 mg, white solid, ¹H NMR (CDCl₃, 400 MHz): δ 7.84-7.80 (m, 2H), 7.75-7.71 (m, 2H), 6.10 (d, *J* = 4 Hz, 1H), 3.17-3.12 (m, 1H), 2.92-2.85 (m, 1H), 2.59 (dd, *J*₁ = 5.6 Hz, *J*₂ = 13.6 Hz, 1H), 2.41-2.35 (m, 1H), 2.27-2.20 (m, 1H), 2.04-1.94 (m, 1H).

¹³C NMR (101 MHz, CDCl₃): δ 163.1, 133.9, 128.4, 123.1, 97.1, 37.4, 32.9, 27.6.

HRMS (ESI) m/z: calcd for C₁₂H₁₁NNaO₃S⁺: M + Na = 272.0352; found: 272.0344.

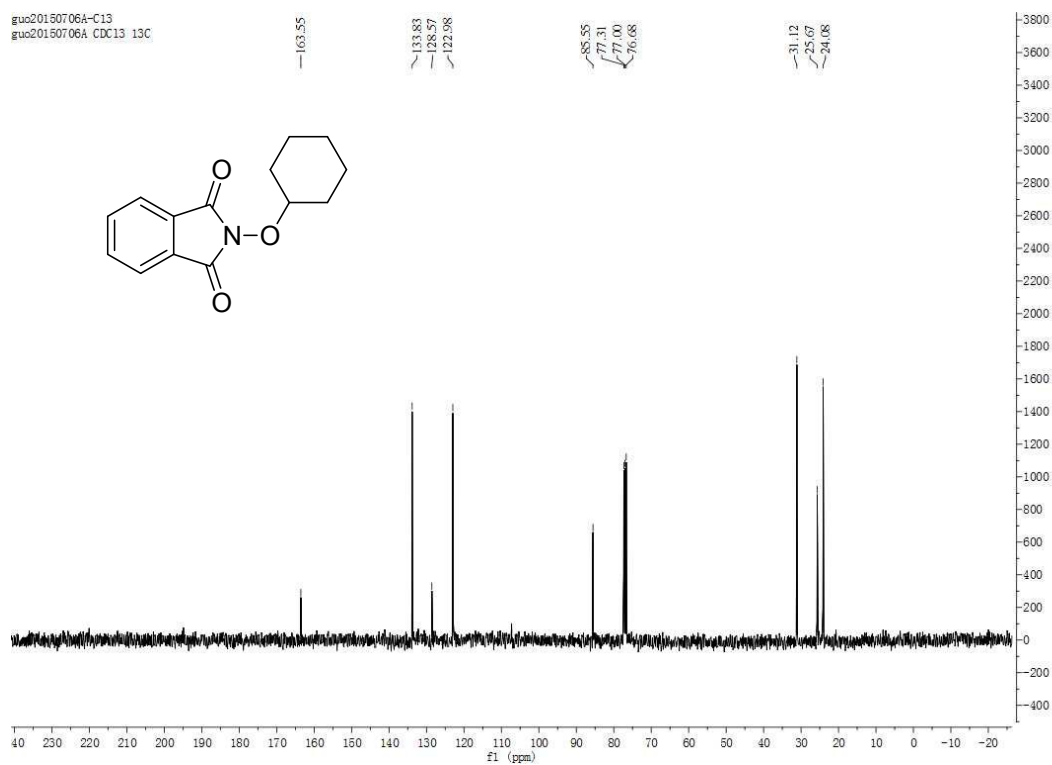
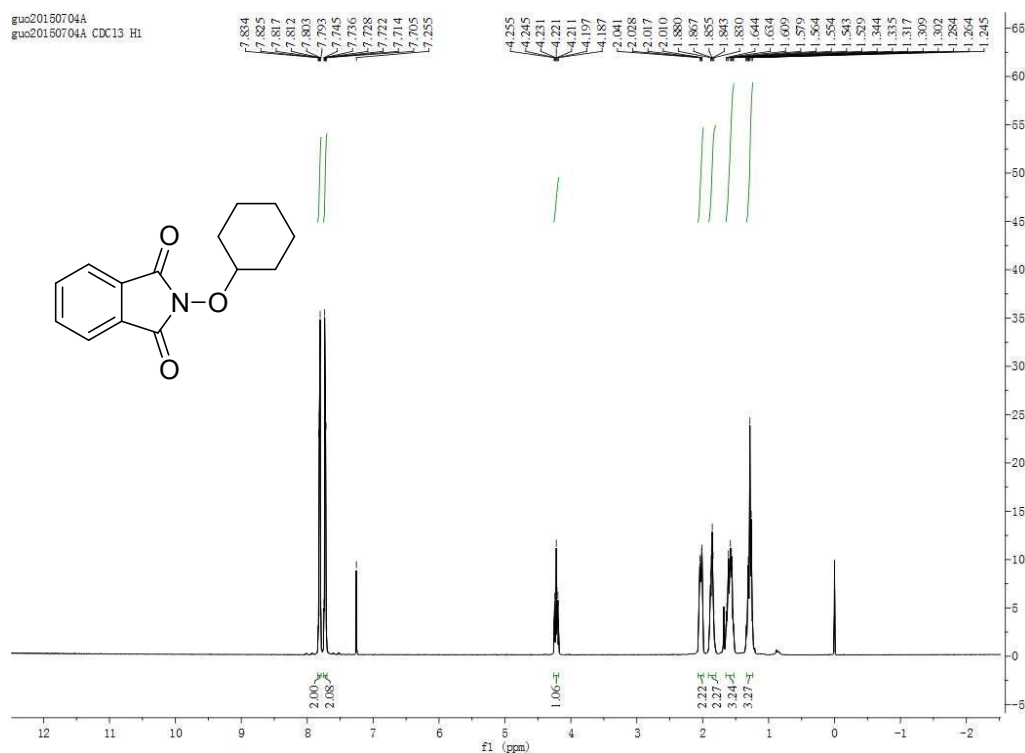
2-(1-(butylthio)butoxy)isoindoline-1,3-dione (Table 2, 2t).

70%, 215 mg, white solid, mp 31-32 °C, ^1H NMR (CDCl_3 , 400 MHz): δ 7.83-7.81 (m, 2H), 7.75-7.23 (m, 2H), 5.36 (t, $J = 6.4$ Hz, 1H), 2.92-2.86 (m, 1H), 2.78-2.71 (m, 1H), 2.14-2.05 (m, 1H), 1.94-1.85 (m, 1H), 1.68-1.51 (m, 4H), 1.45-1.36 (m, 2H), 1.00 (t, $J = 7.2$ Hz, 3H), 0.91 (t, $J = 7.2$ Hz, 3H).

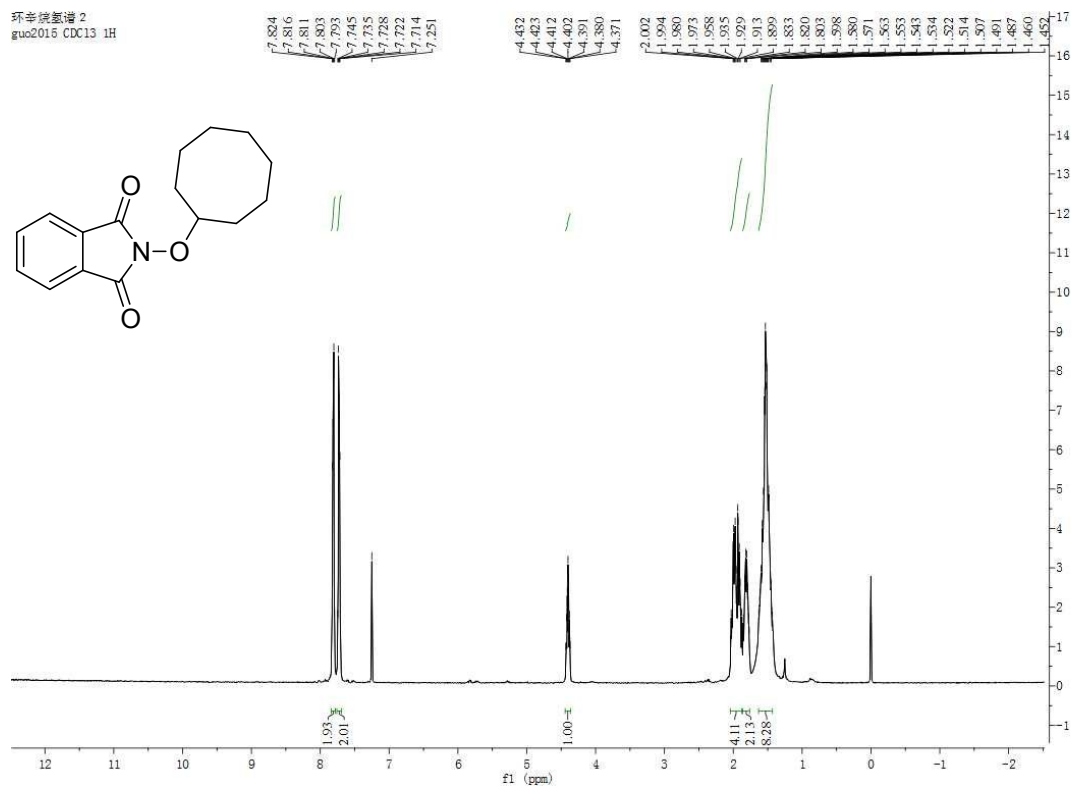
^{13}C NMR (101 MHz, CDCl_3): δ 162.6, 133.5, 128.2, 122.5, 93.2, 35.6, 31.7, 28.6, 22.2, 19.6, 13.8.

HRMS (ESI) m/z : calcd for $\text{C}_{16}\text{H}_{21}\text{NNaO}_3\text{S}^+$: $M + \text{Na} = 330.1134$; found: 330.1119.

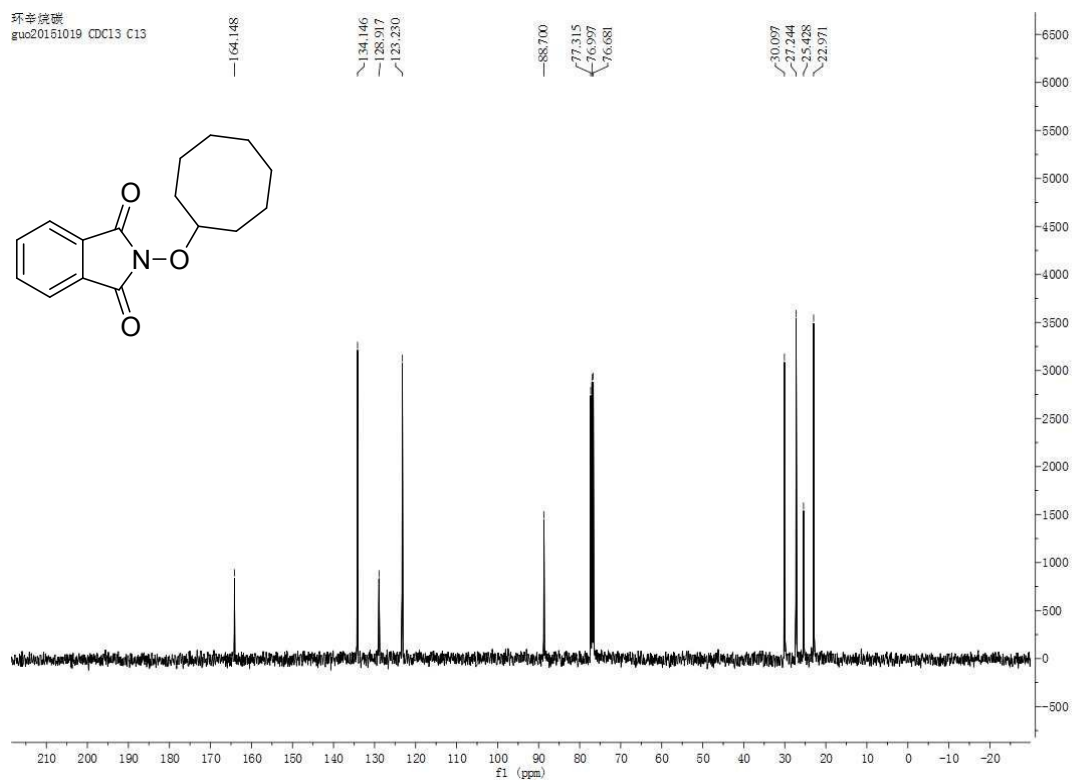
NMR spectra of product



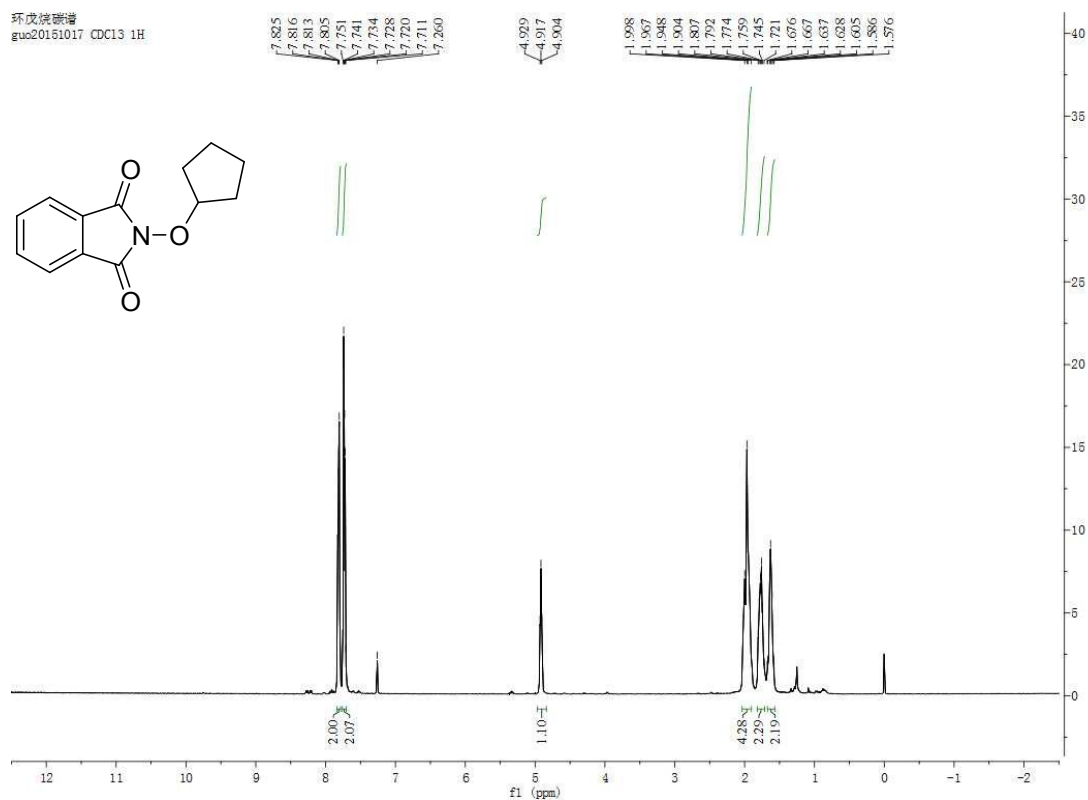
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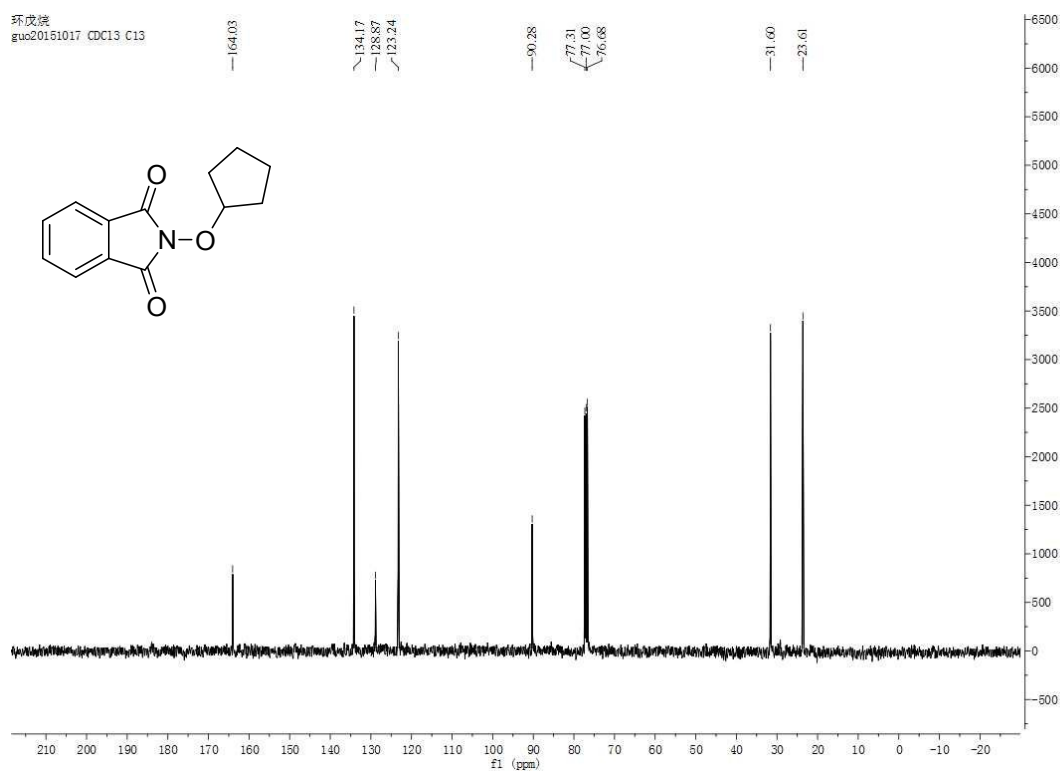
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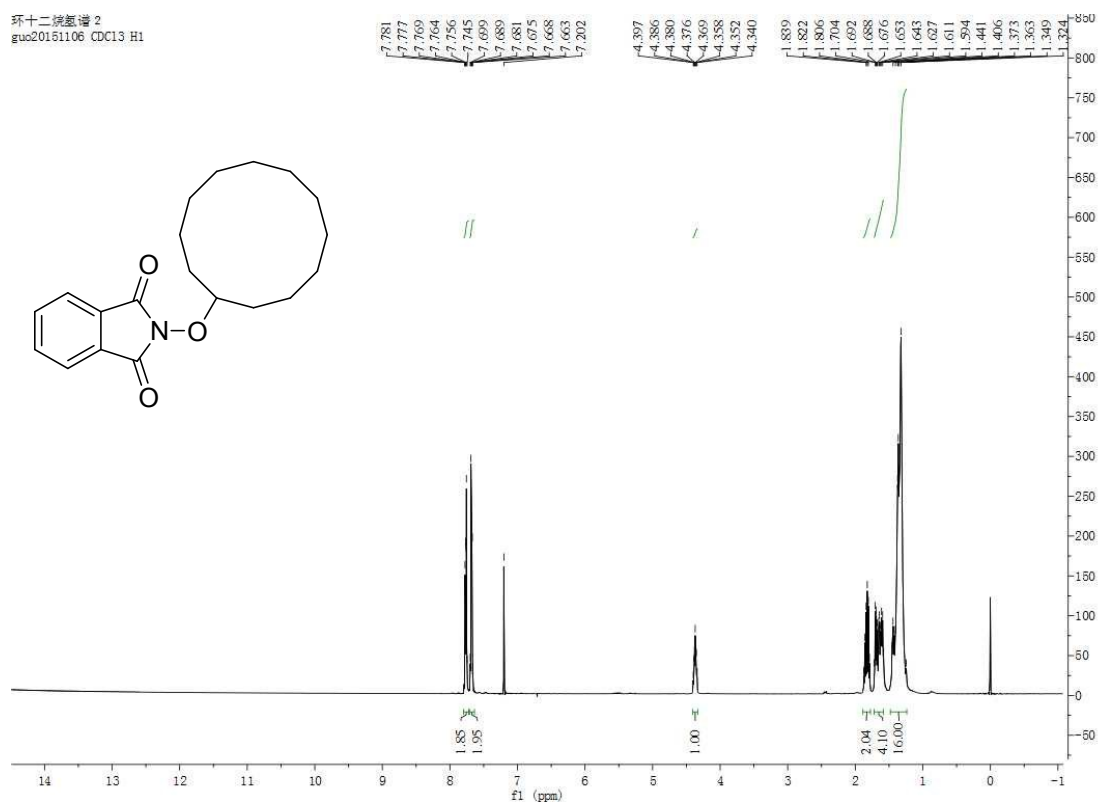
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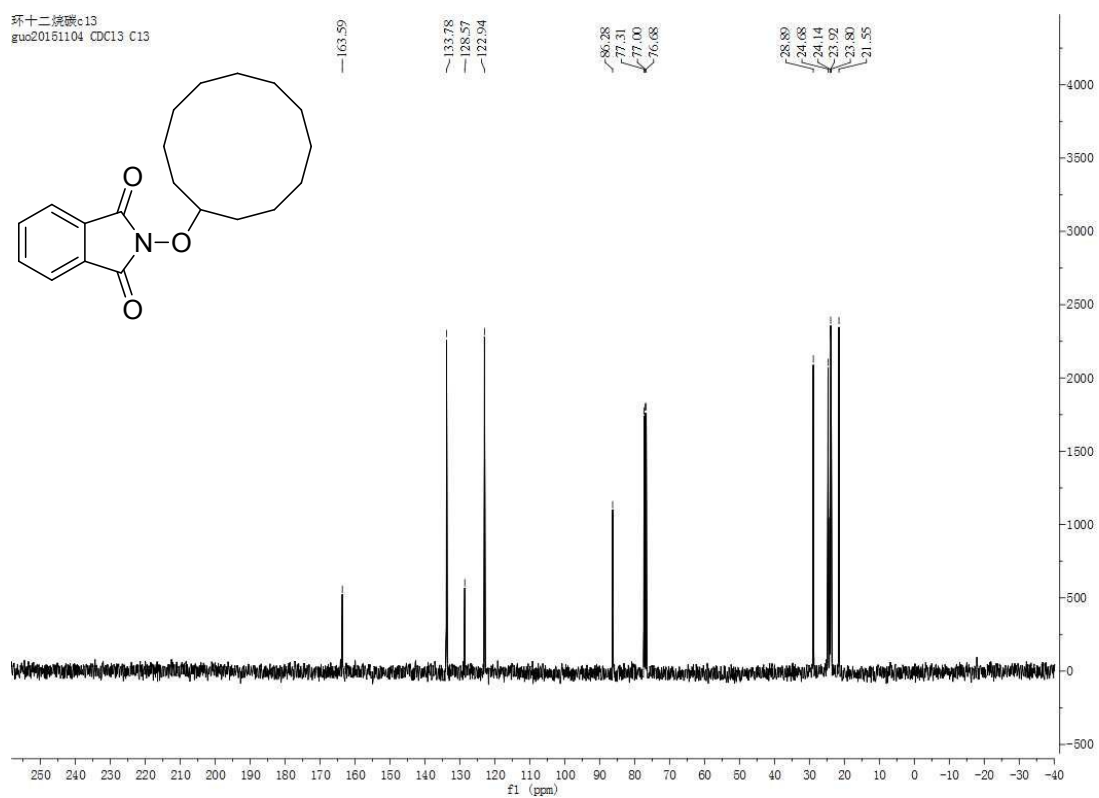
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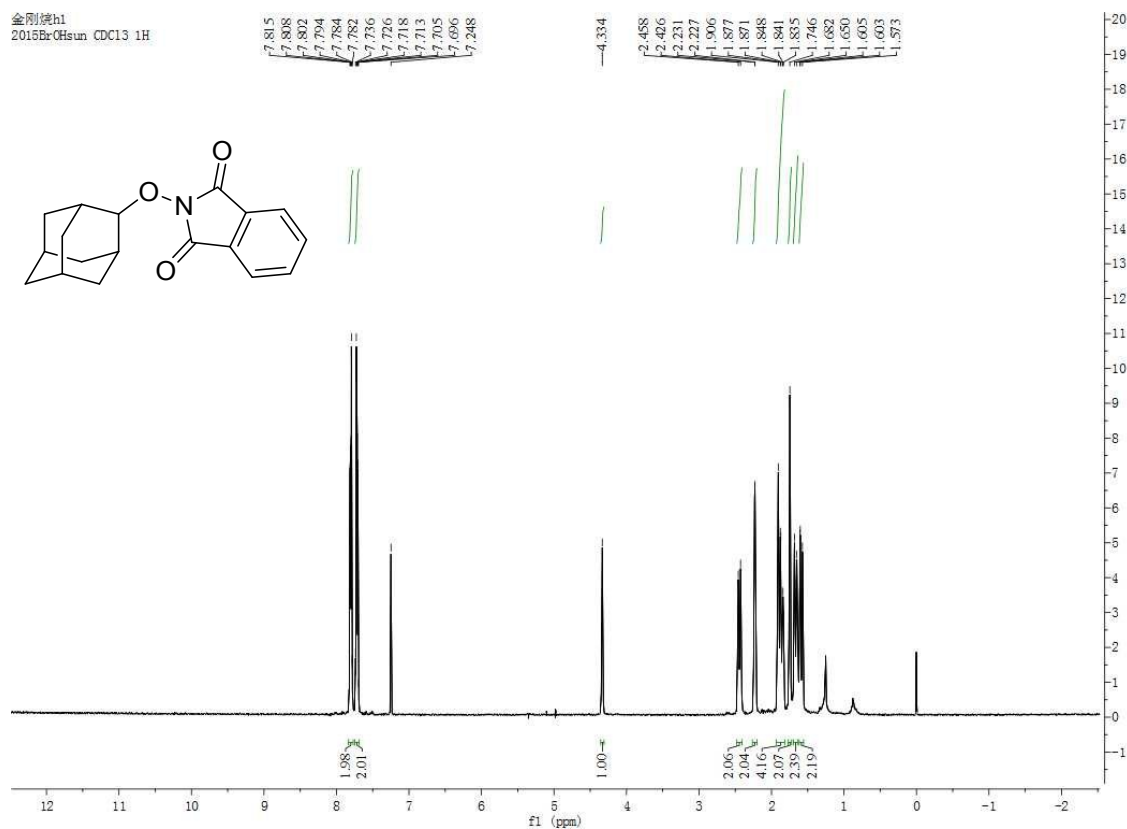
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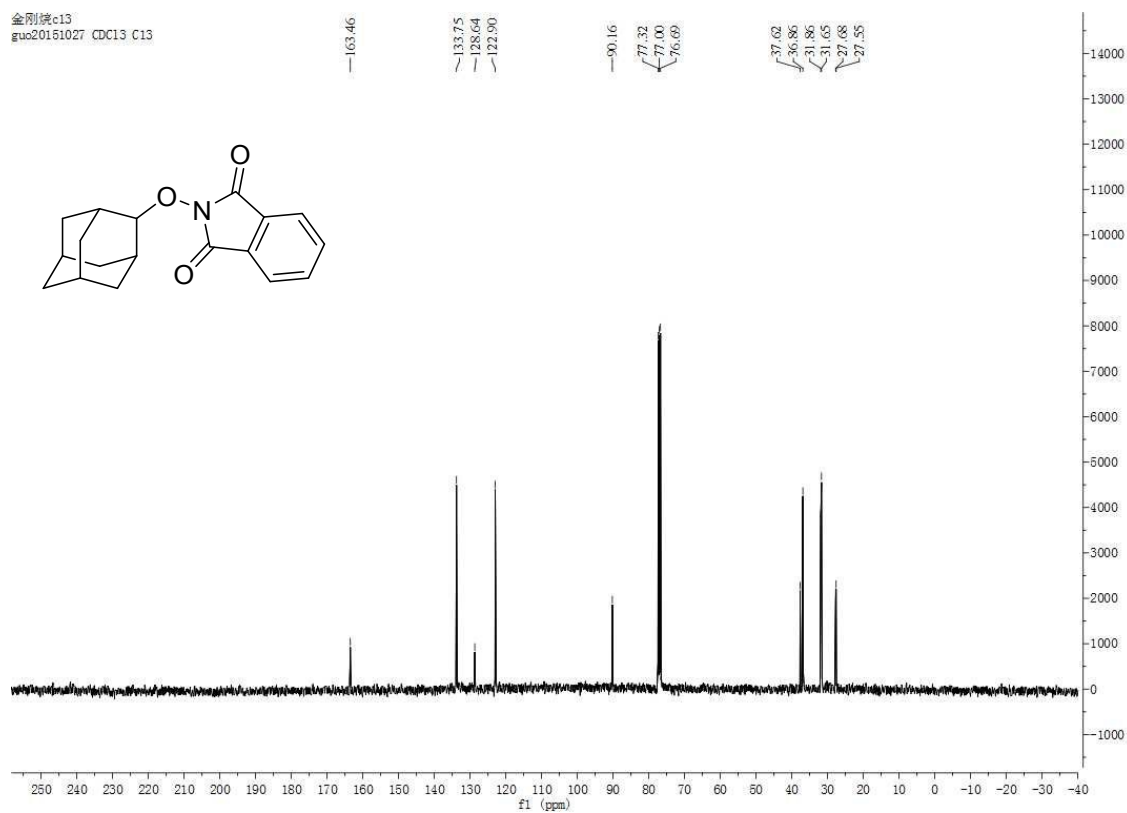
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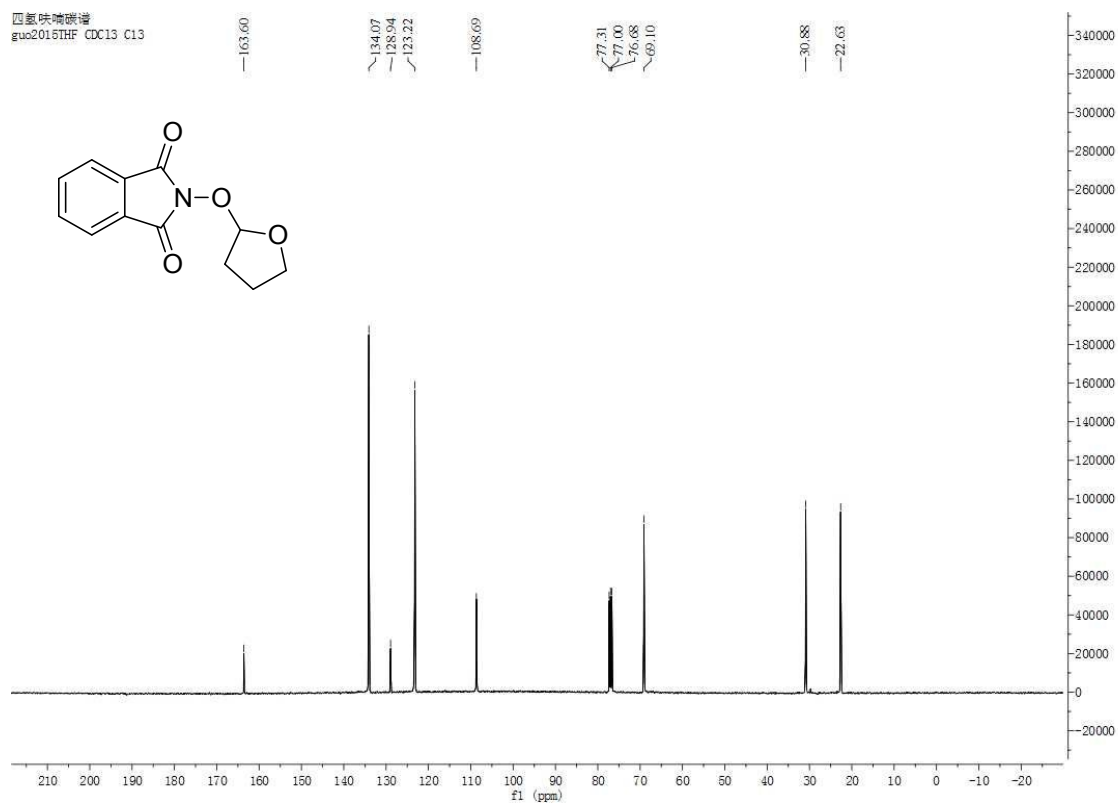
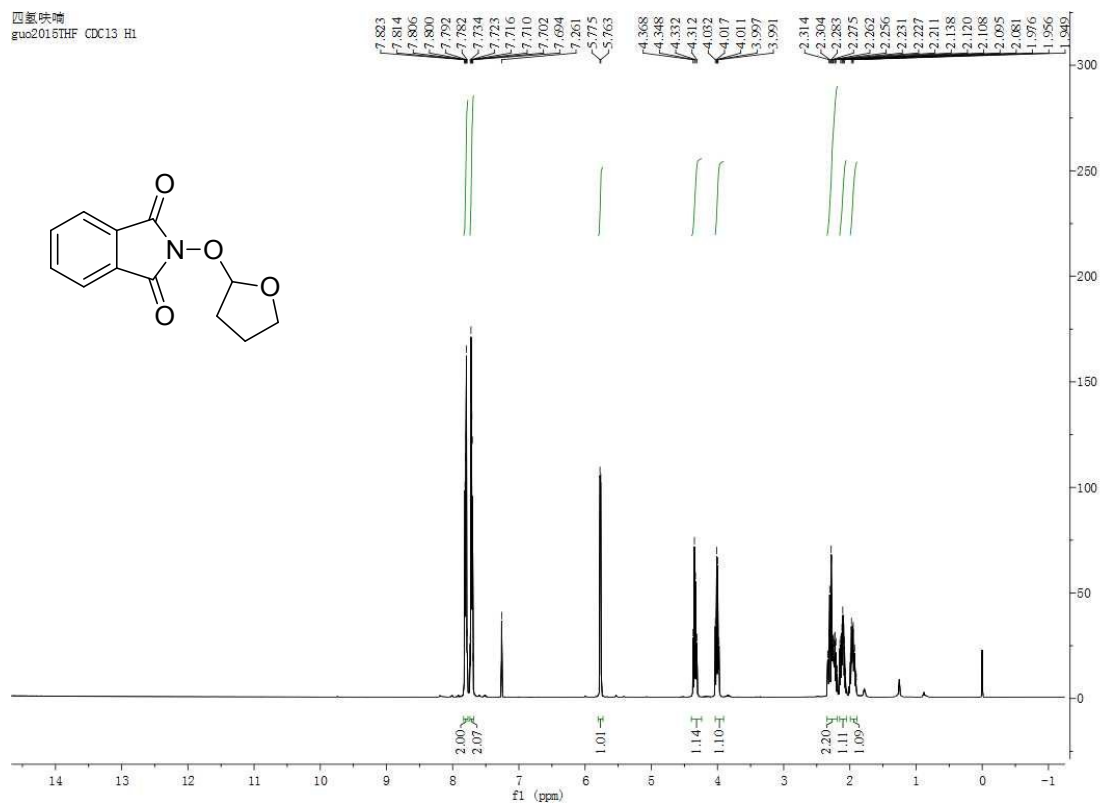


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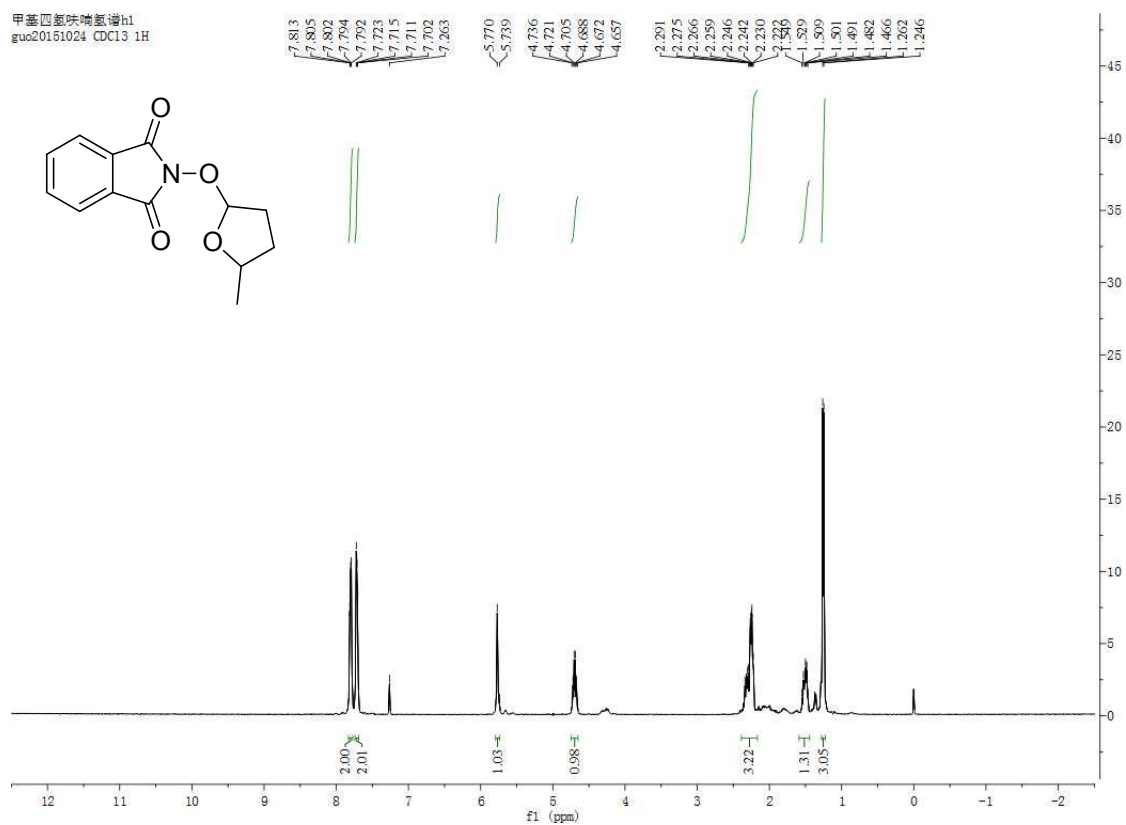


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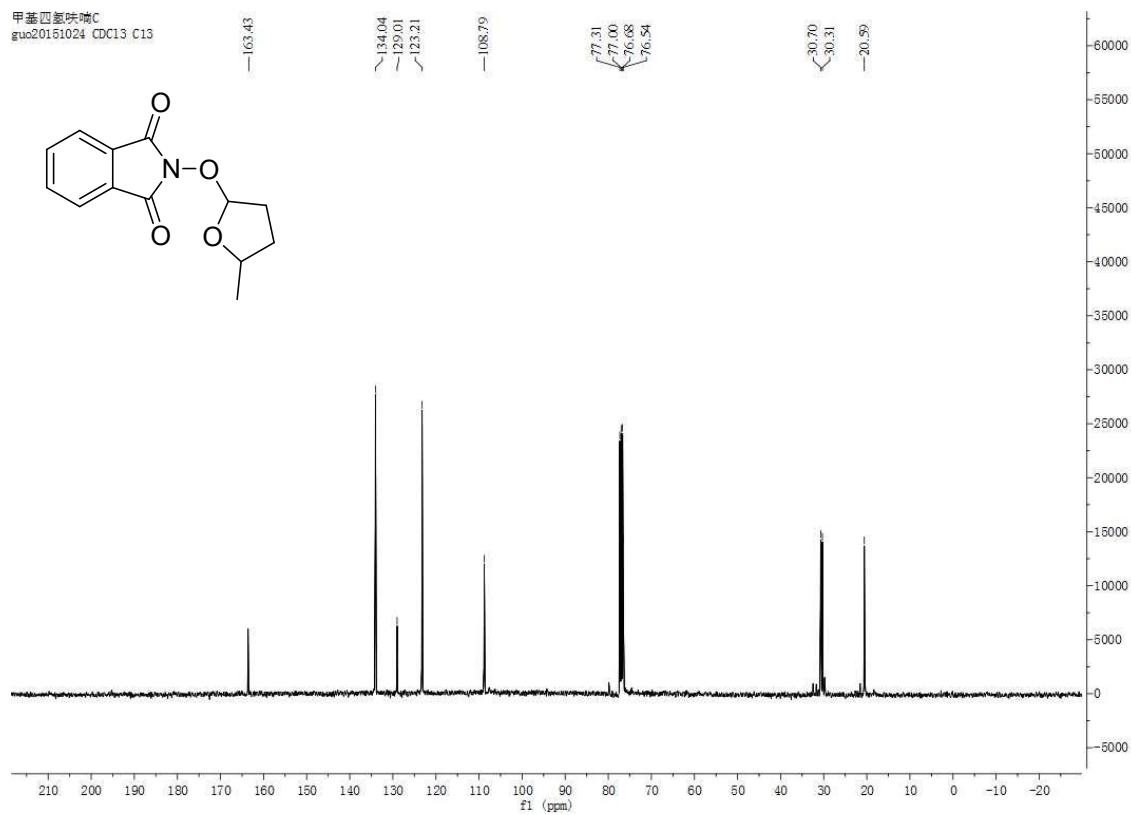




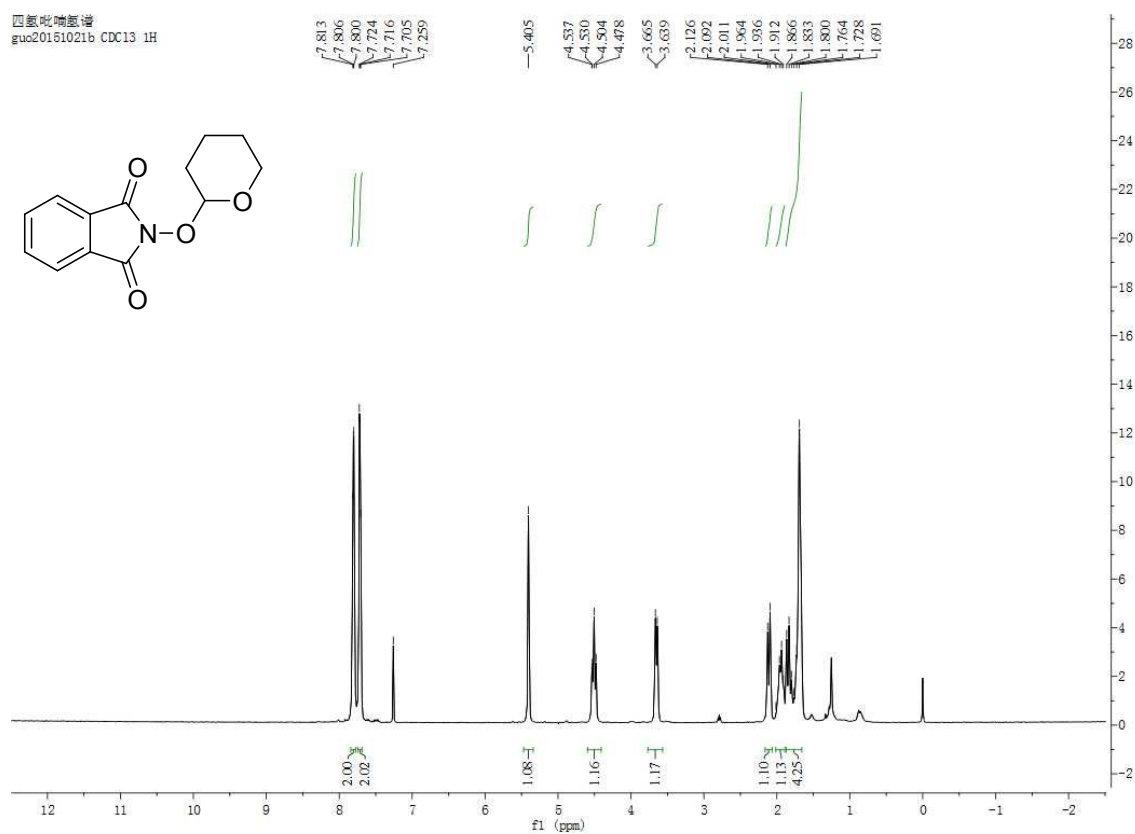
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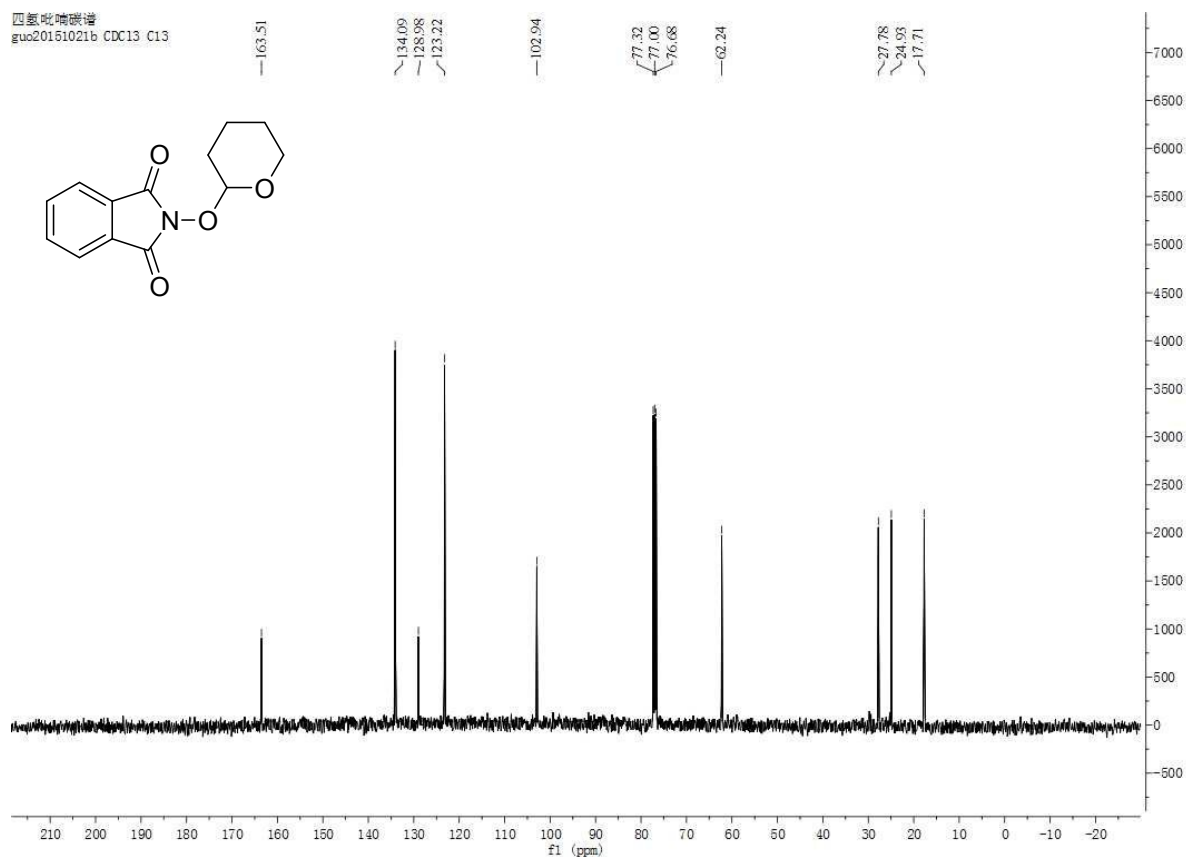
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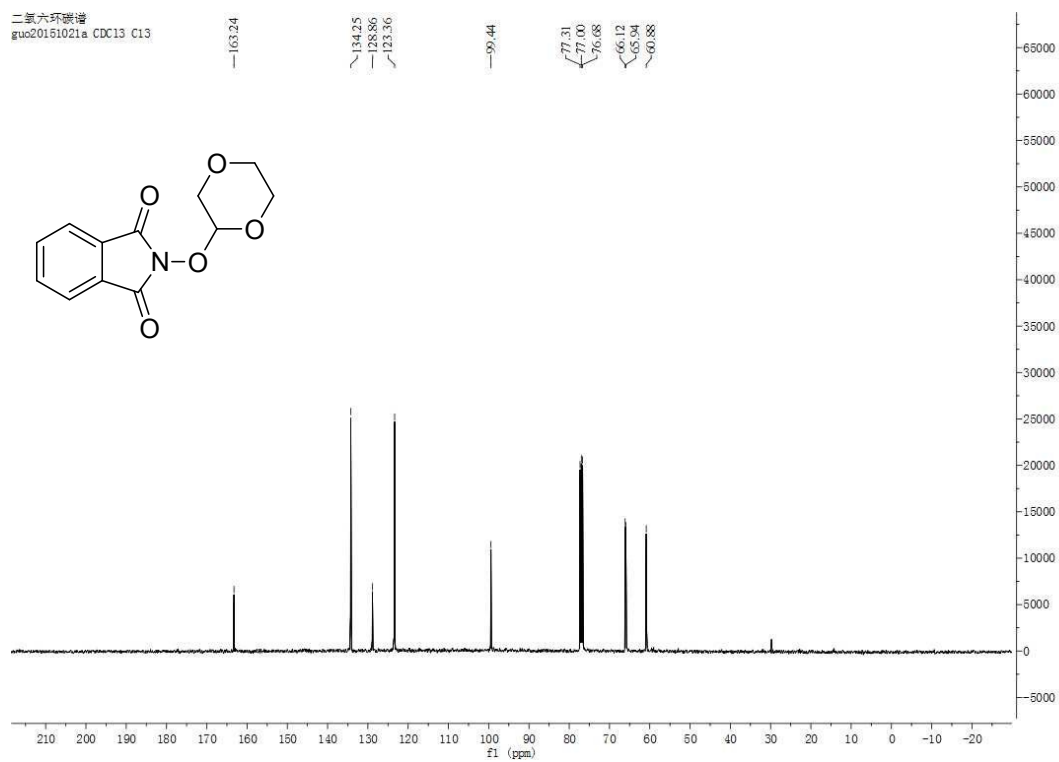
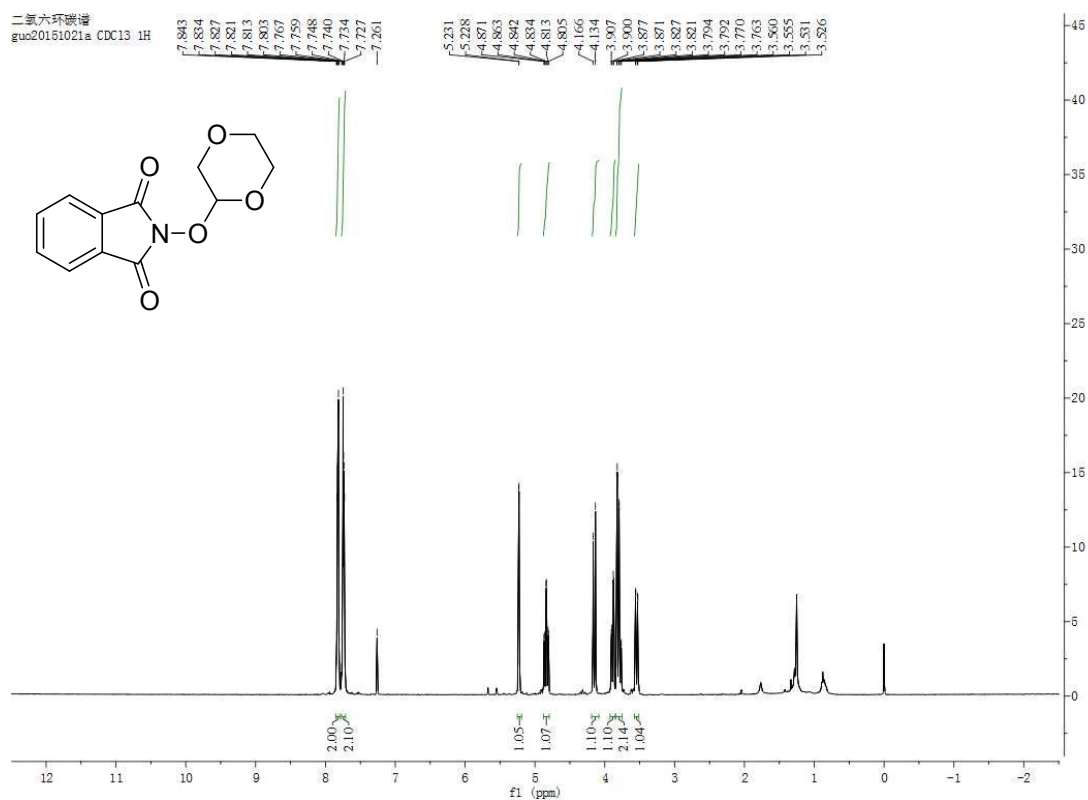


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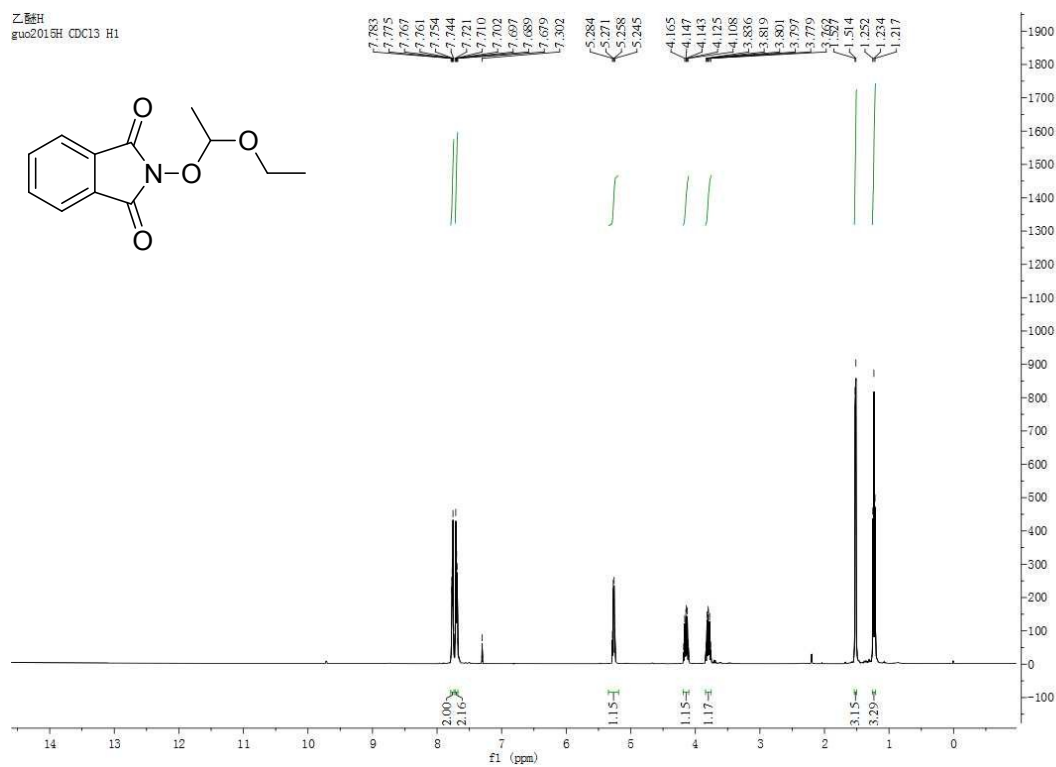


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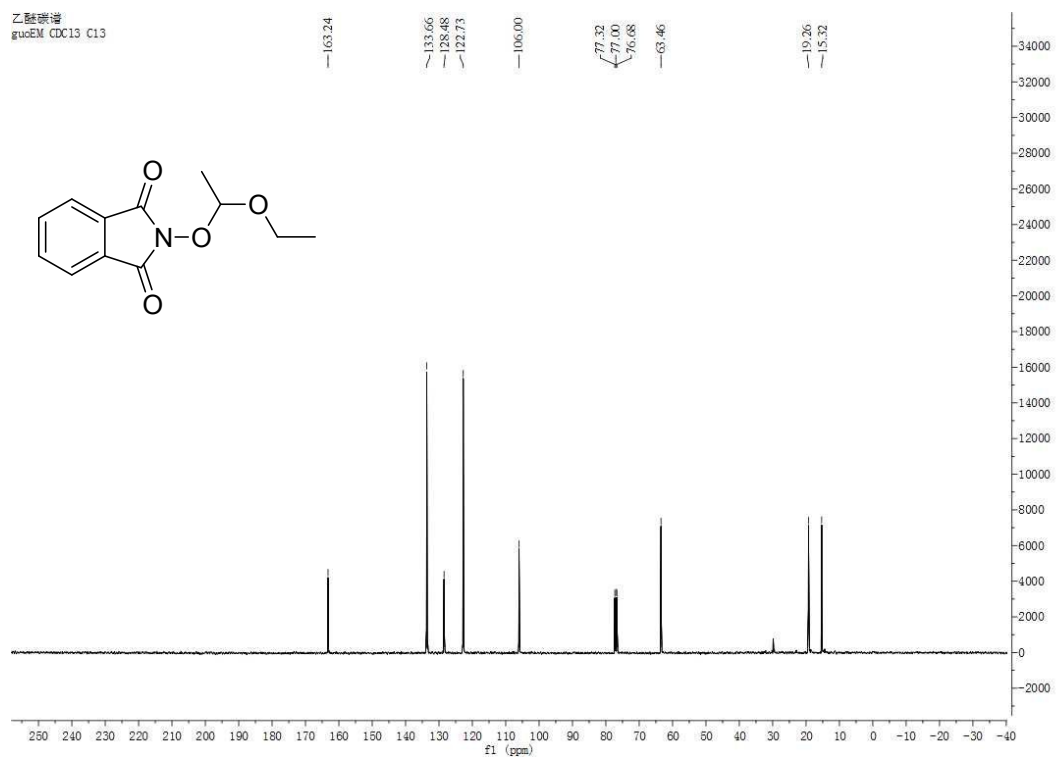


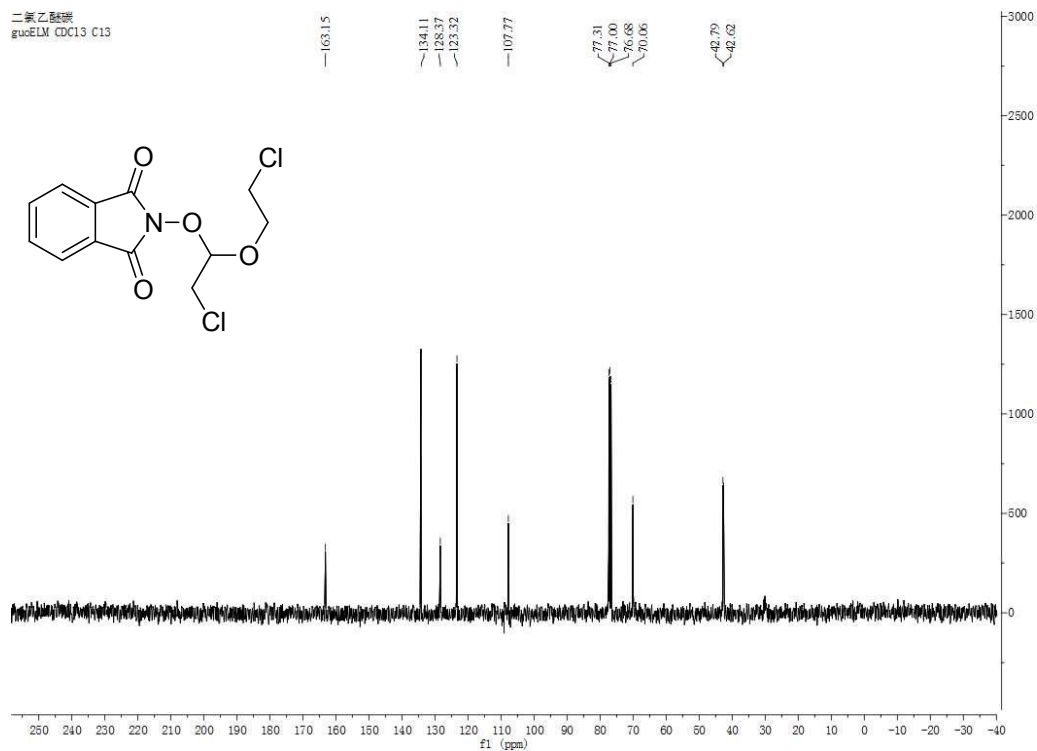
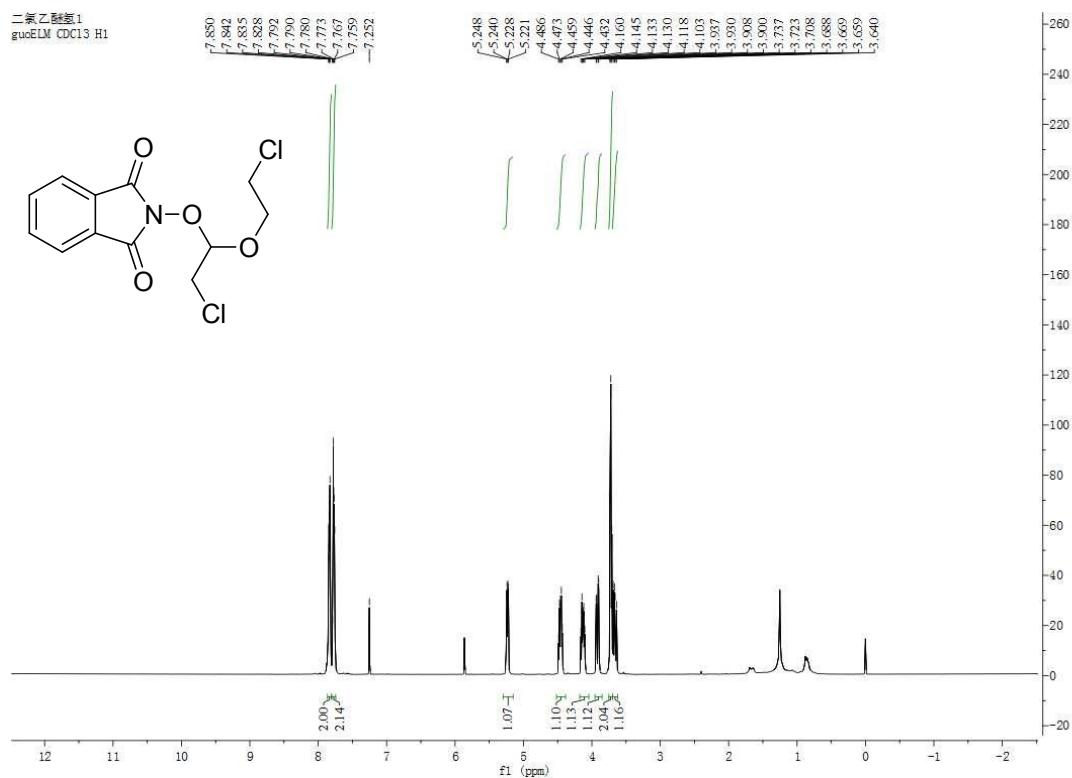


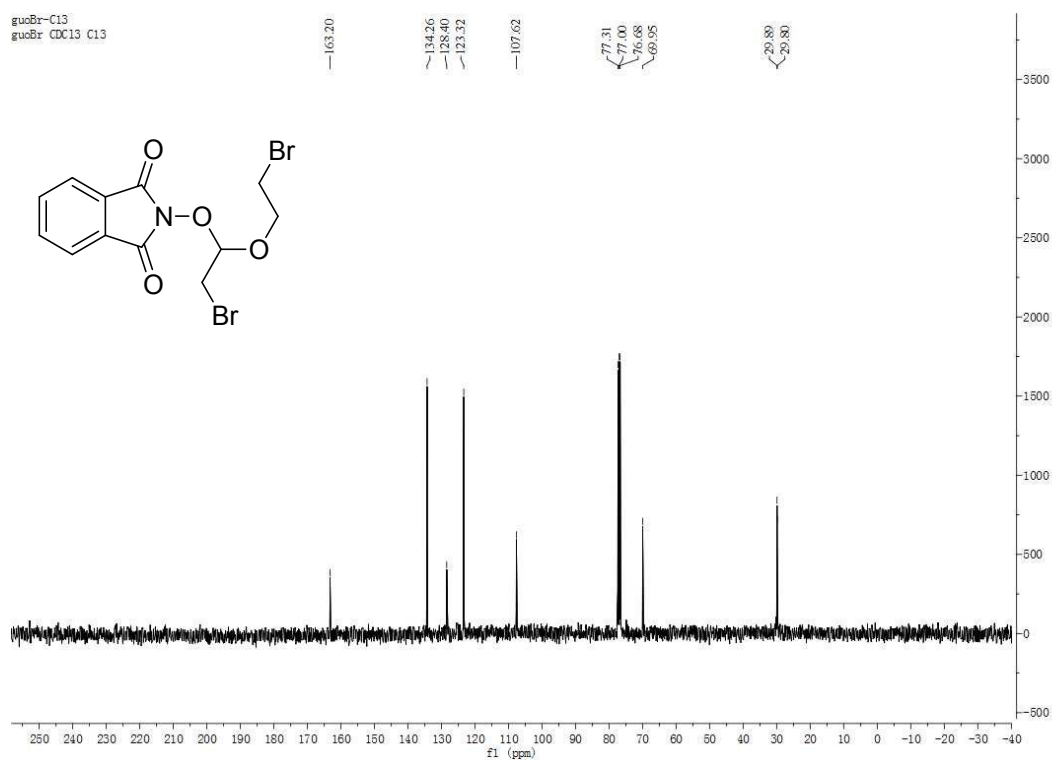
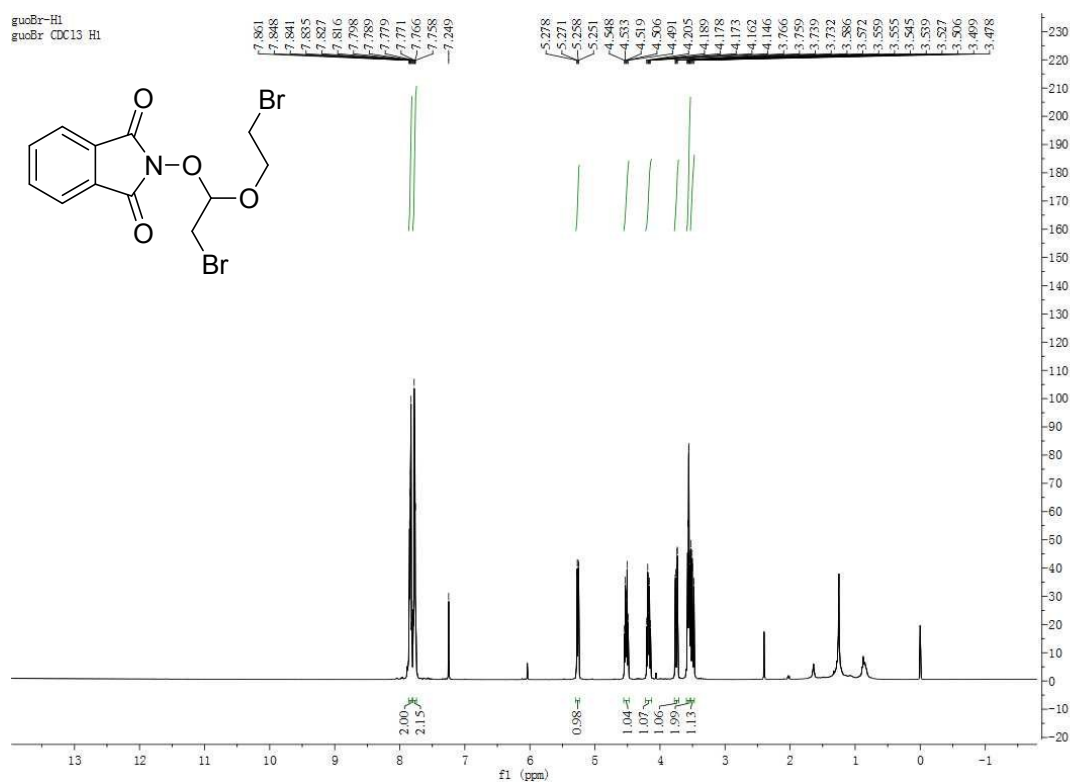
乙酰基
guo2018H CDC13 H1

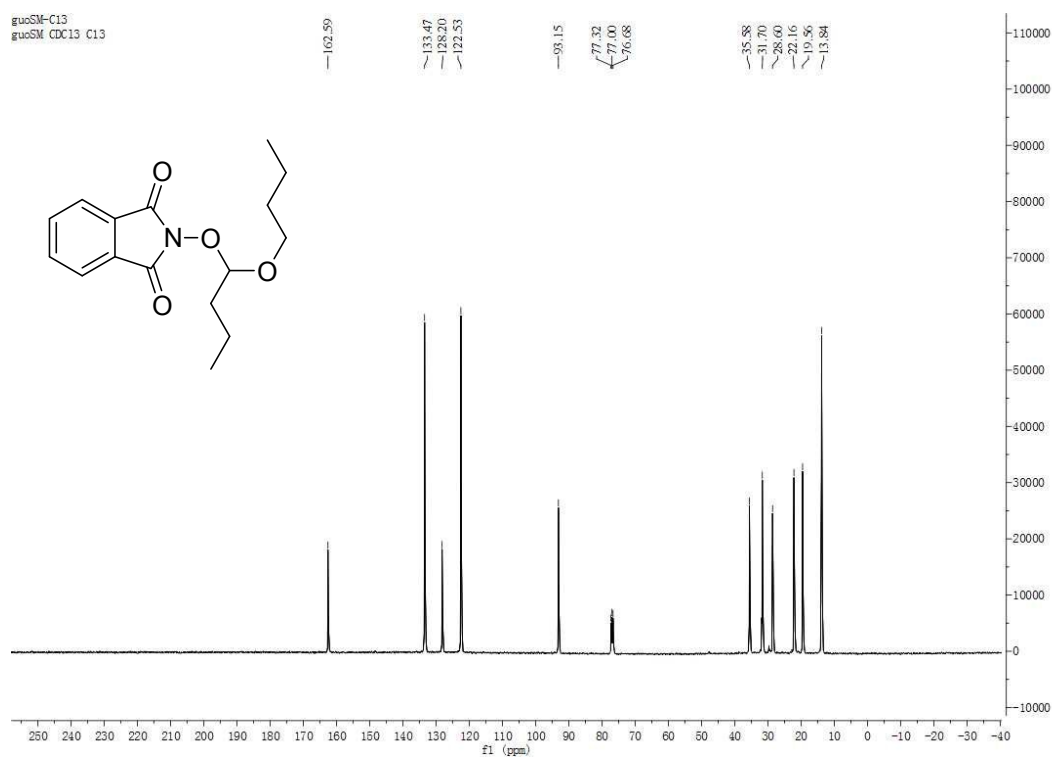
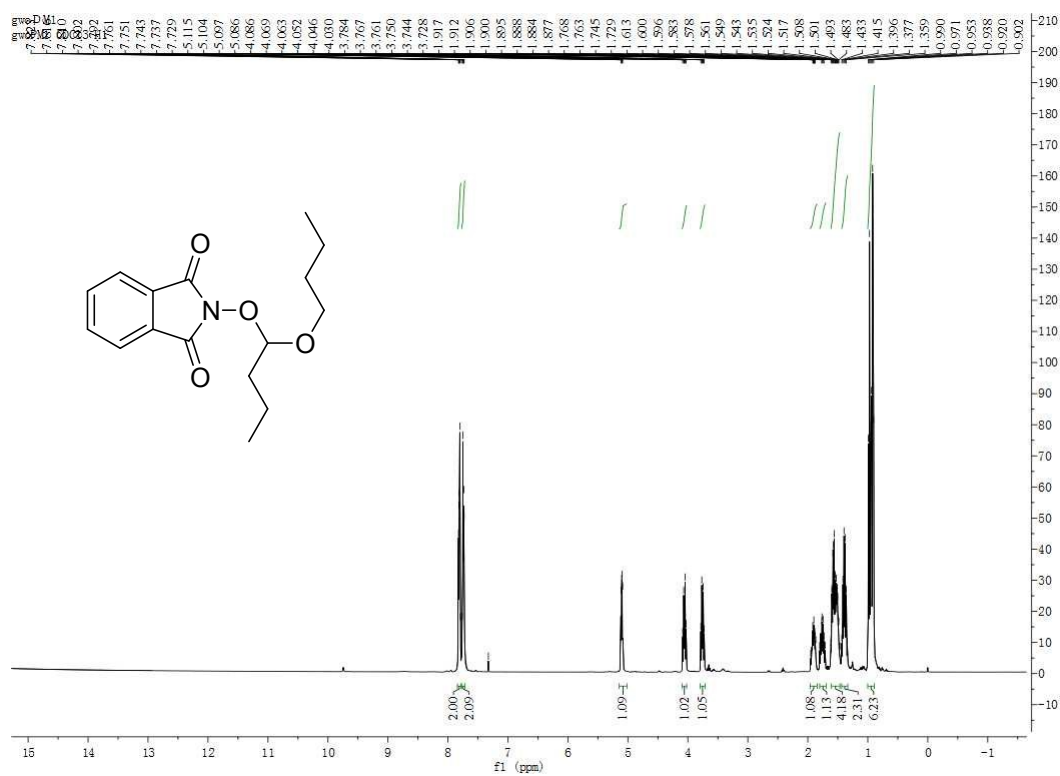


乙酰基
guoEM CDC13 C13

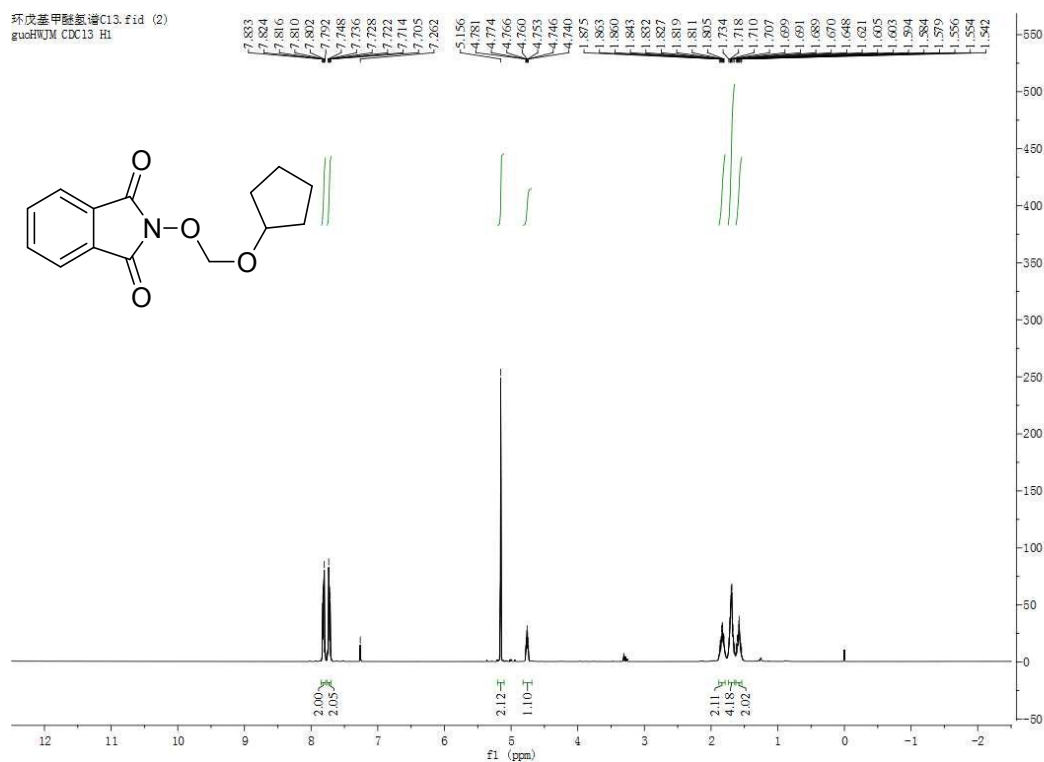




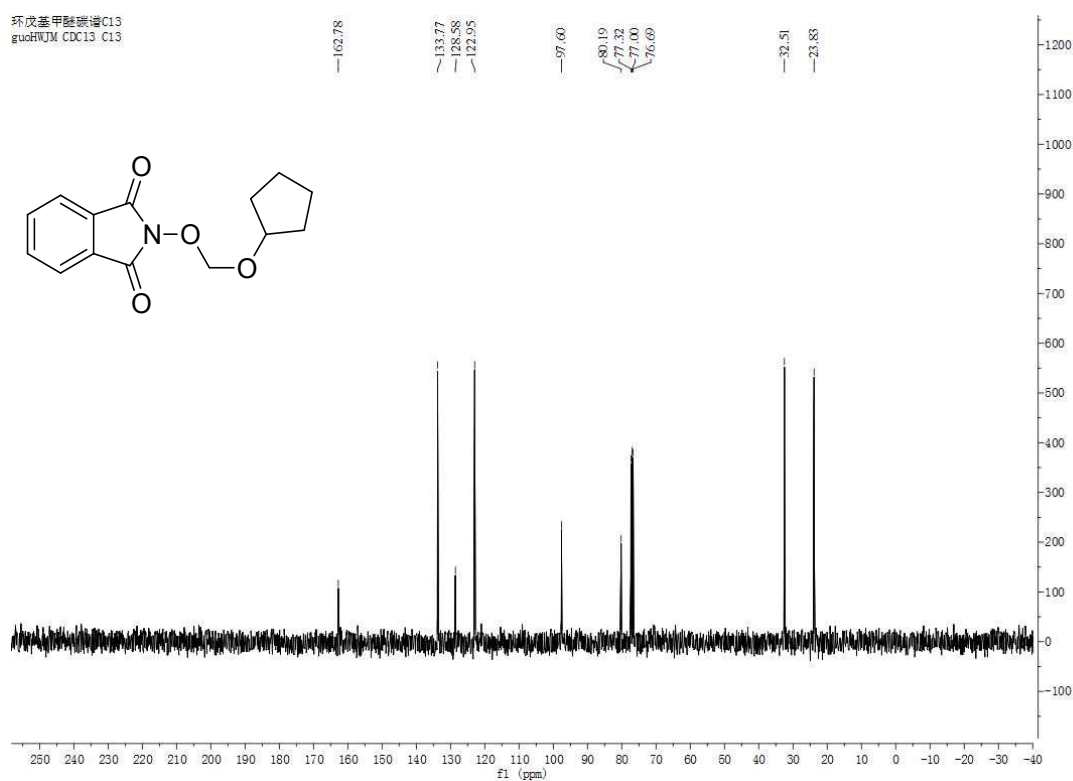




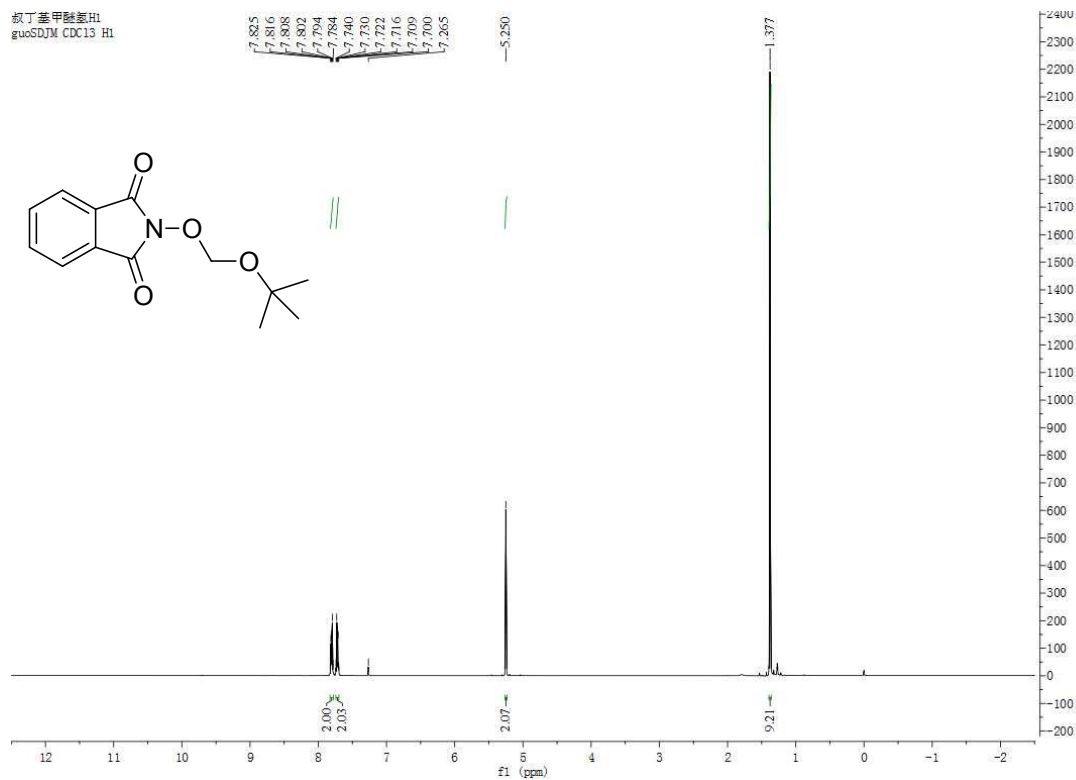
环戊基甲醚谱C13.fid (2)
guohuim CDC13 H1



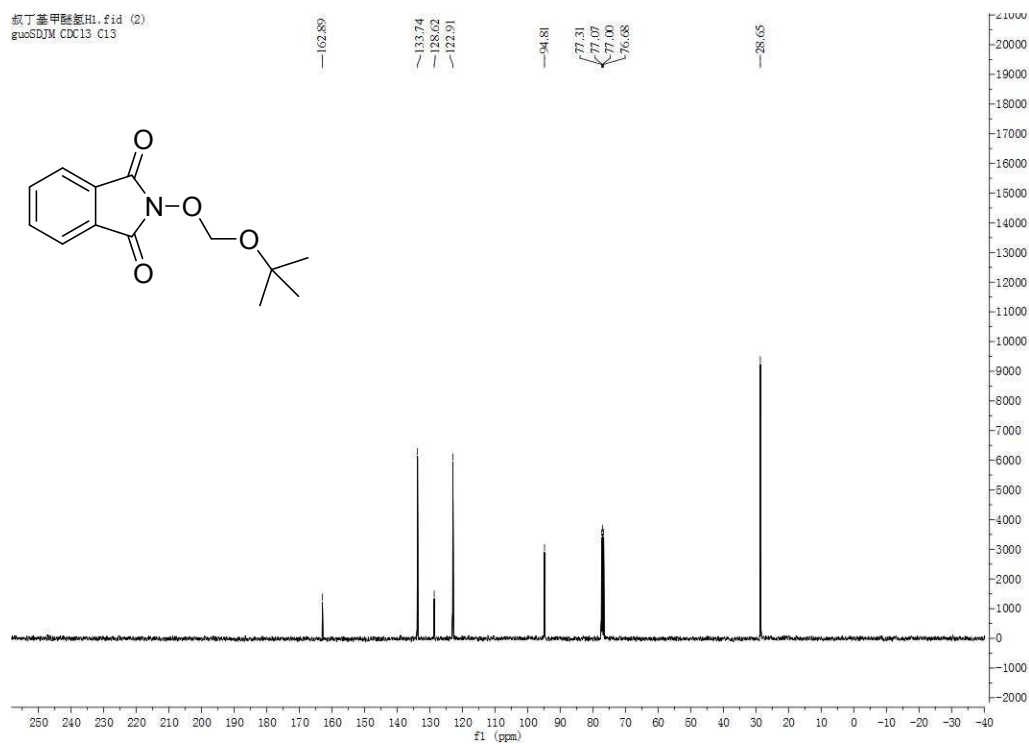
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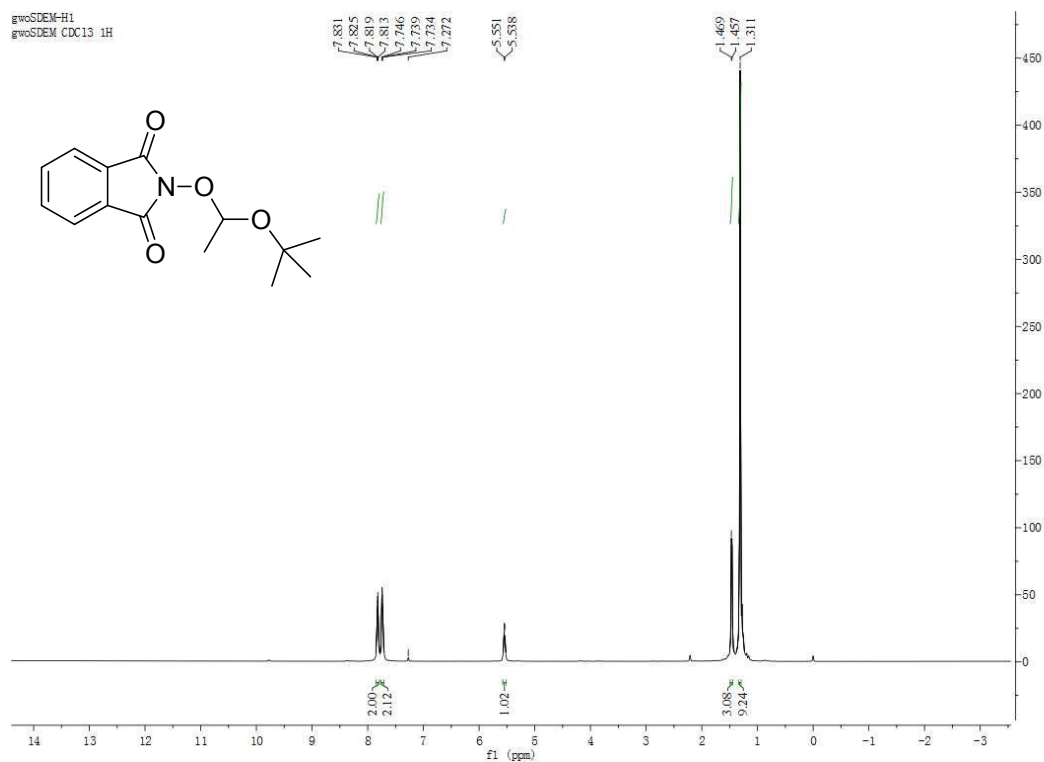
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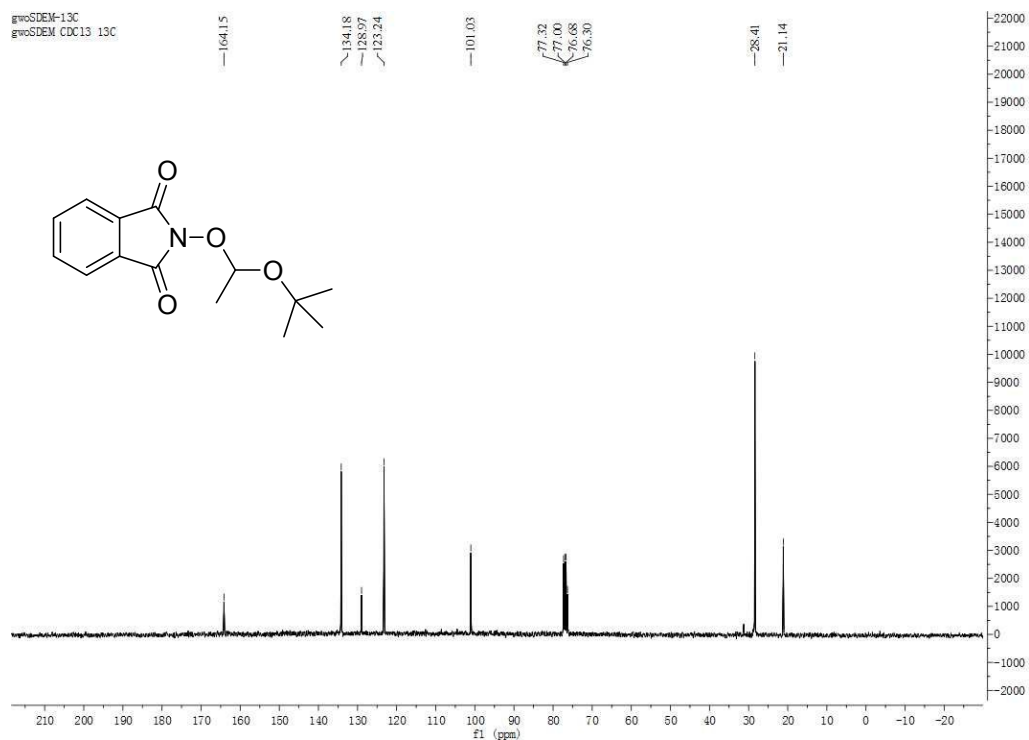
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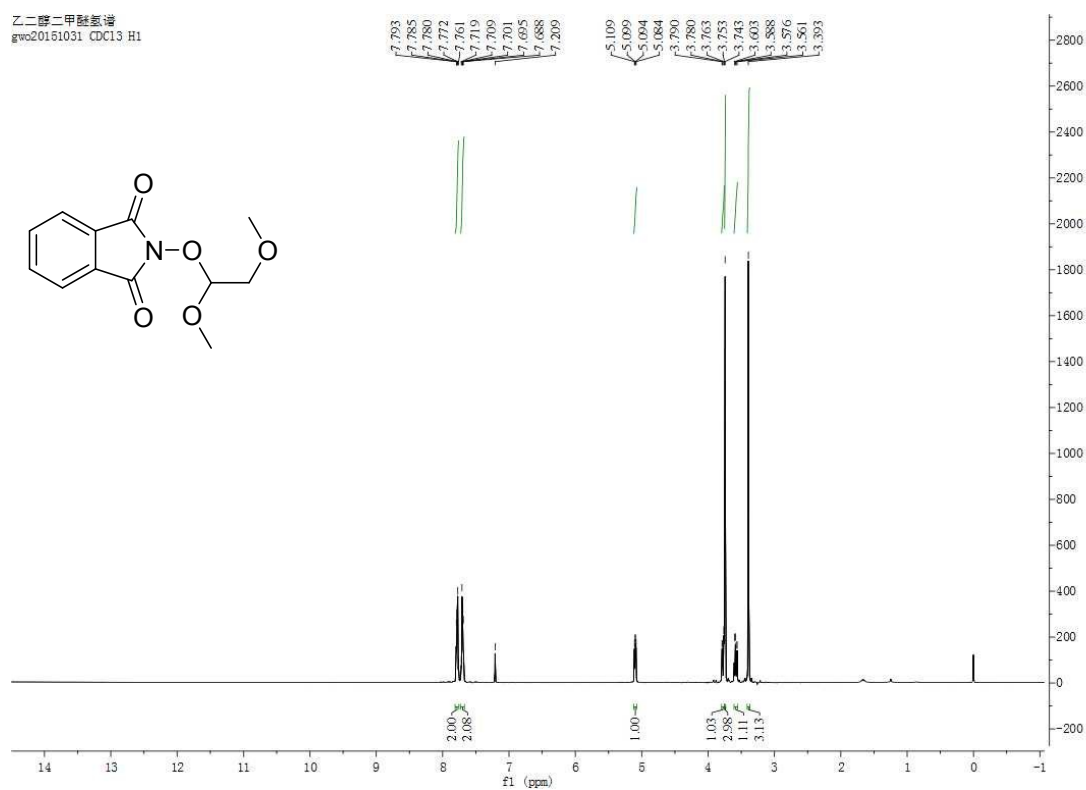
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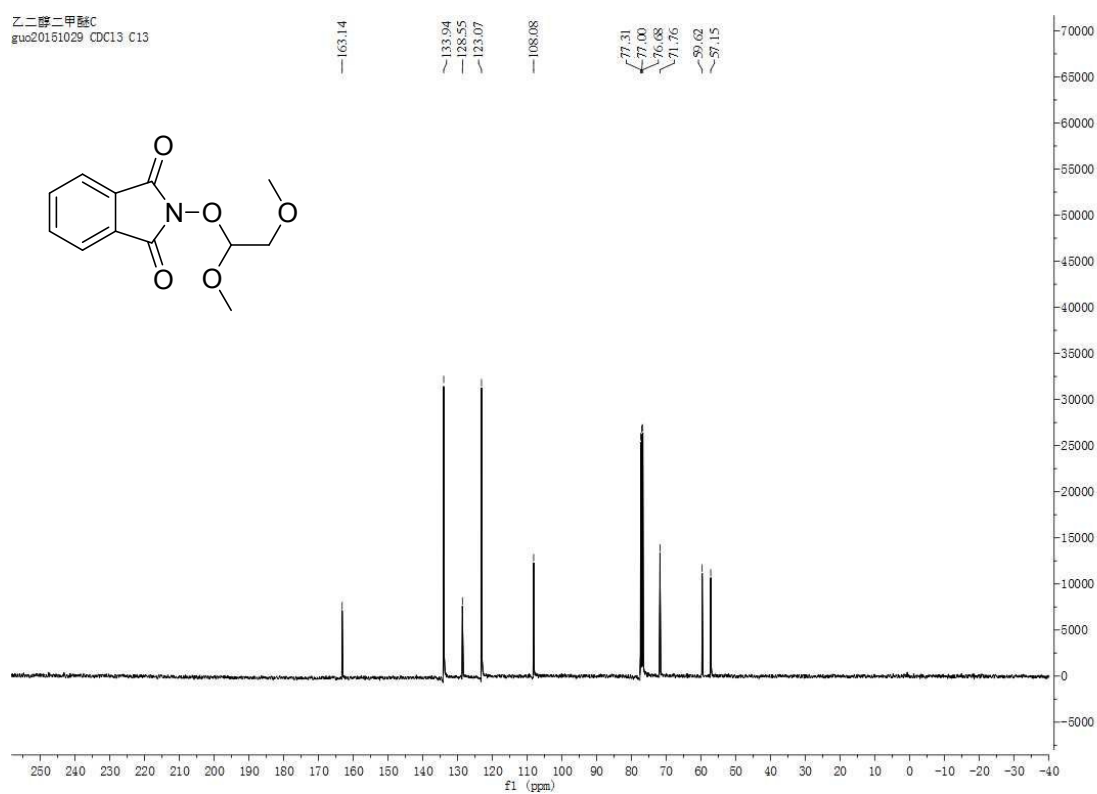
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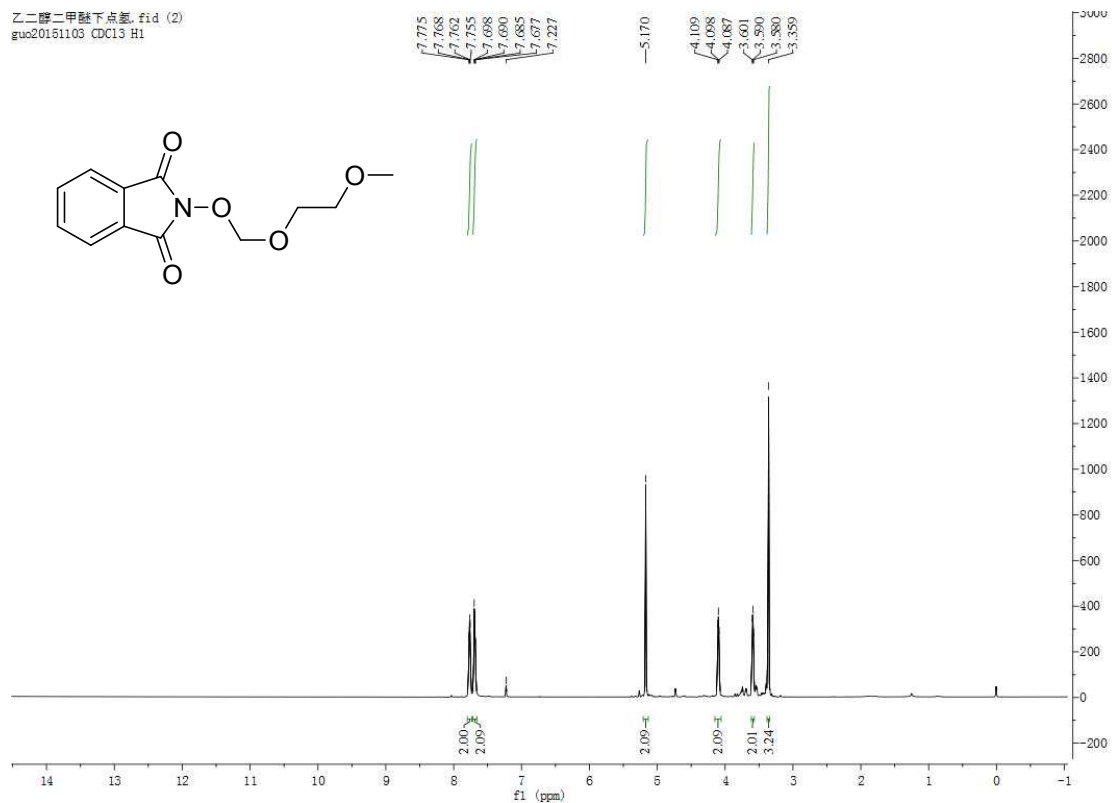
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guo20151031 CDC13 H1



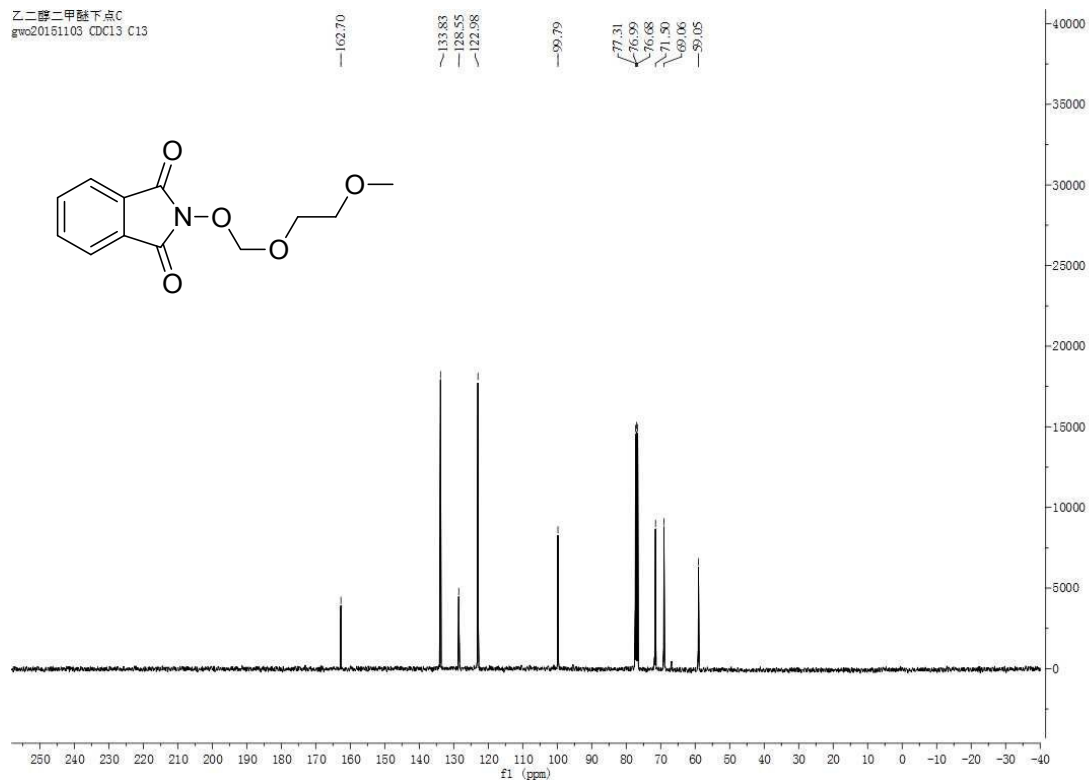
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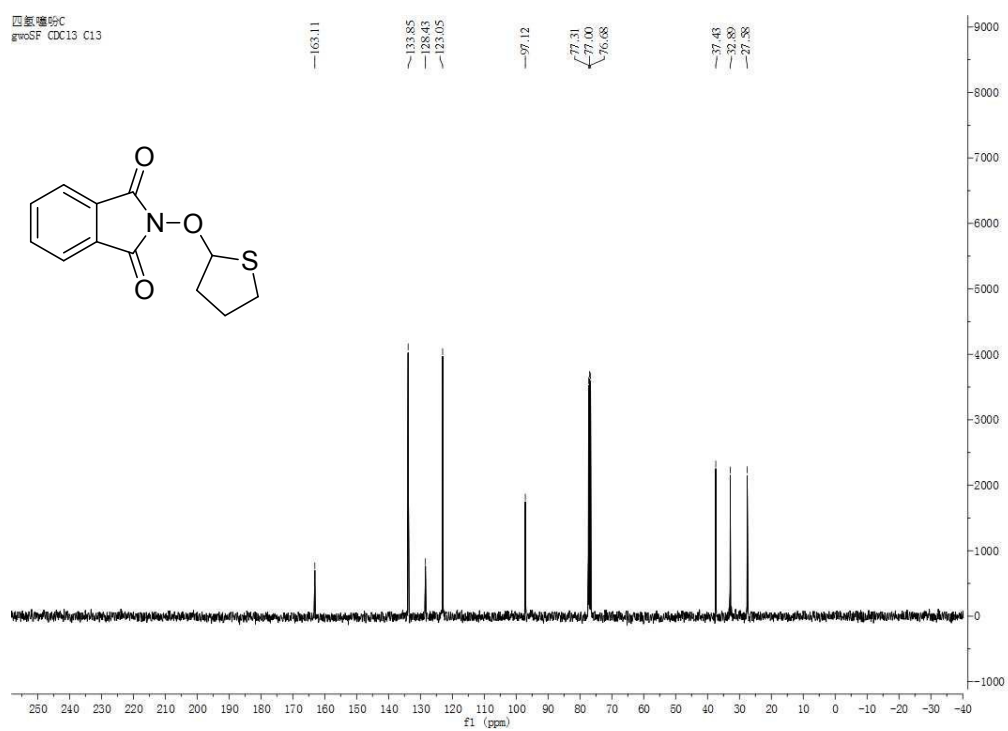
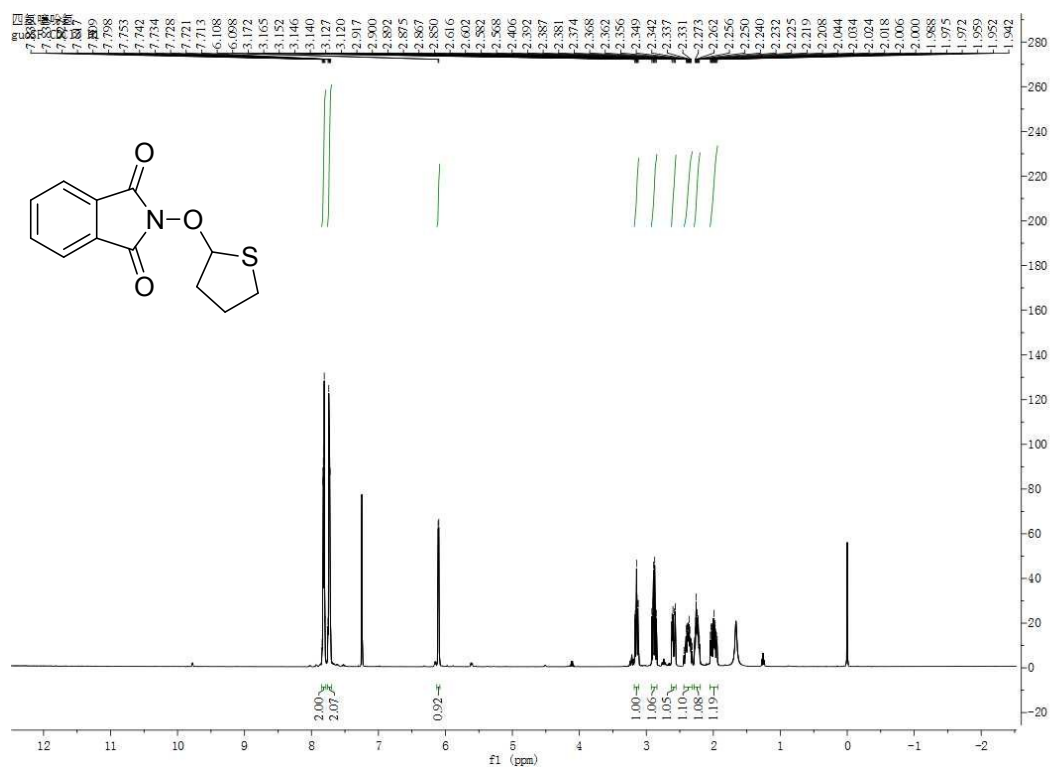


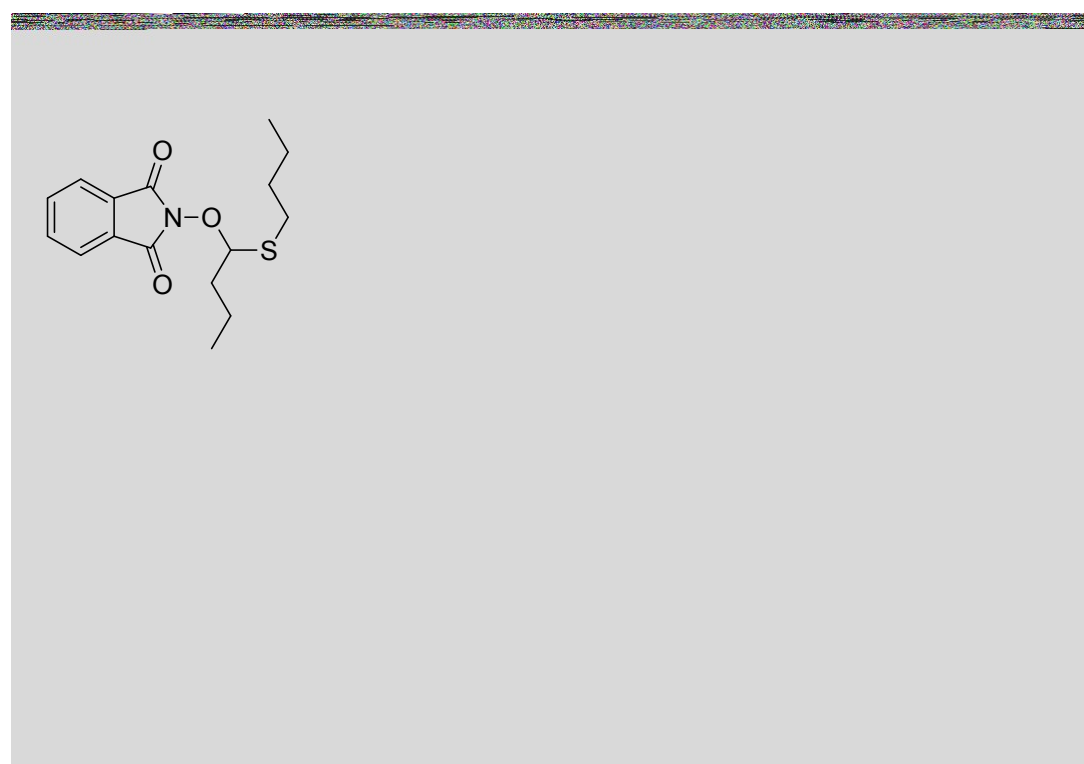
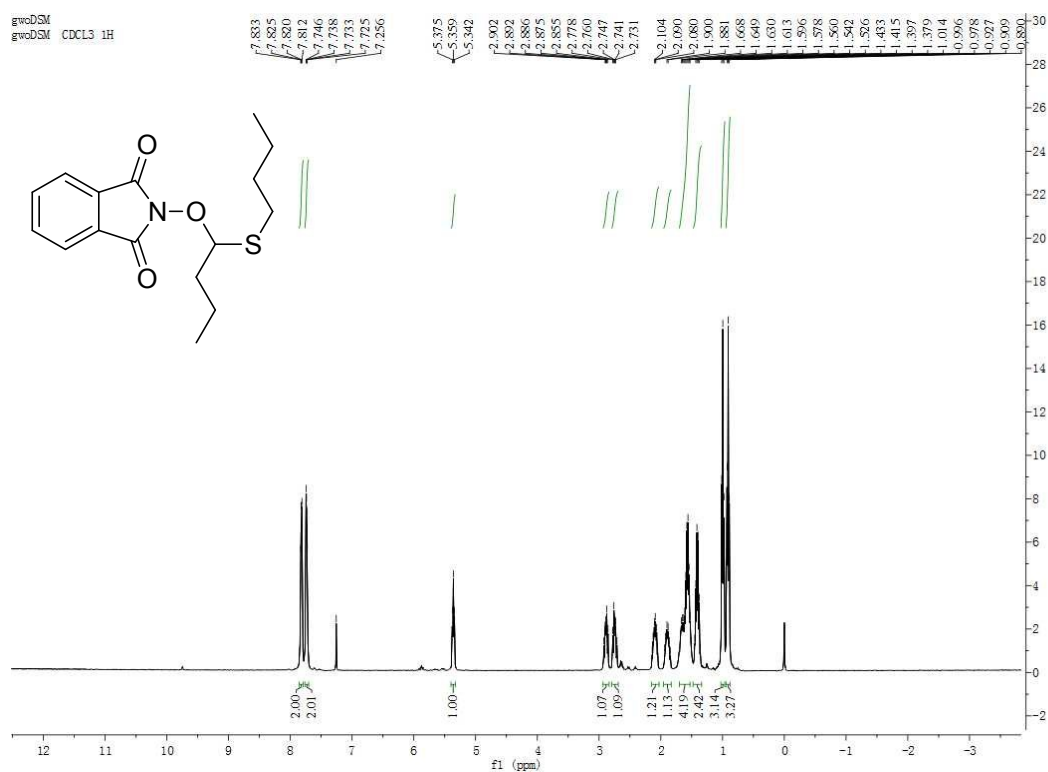
乙二醇二甲醚下点氮, fid (2)
guo20151103 CDC13 H1



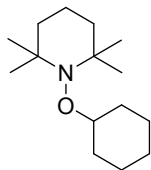
乙二醇二甲醚下点C
guo20151103 CDC13 C13







Electrospray ionization (ESI) spectra of 1-(cyclohexyloxy)-2,2,6,6-tetramethylpiperidine



1-(cyclohexyloxy)-2,2,6,6-tetramethylpiperidine

Chemical Formula: C₁₅H₂₉NO

Exact Mass: 239.22

(ESI) m/z: M + H = 240.3

