

Supporting Information

for

**Three-component synthesis, utilization and biological activity of
phosphinoyl-functionalized isoindolinones**

Nóra Popovics-Tóth,^a Bettina Rávai,^a Ádám Tajti,^a Bence Varga,^a Péter Bagi,^a Franc Perdih,^b
Pál Tamás Szabó,^c László Hackler Jr.,^d László G. Puskás^d and Erika Bálint*^a

^a*Department of Organic Chemistry and Technology, Budapest University of Technology and
Economics, 1521 Budapest, Hungary*

^b*Faculty of Chemistry and Chemical Technology, University of Ljubljana, SI-1000 Ljubljana,
Slovenia*

^c*MS Metabolomics Research Group, Centre for Structural Study, Research Centre for Natural
Sciences, Eötvös Loránd Research Network, Magyar tudósok krt. 2, H-1117 Budapest,
Hungary*

^d*Avidin Ltd., Alsó kikötő sor 11/D, H-6726 Szeged, Hungary*

Table of contents

General information.....	S3
General procedure for the synthesis of 3-oxoisoindolin-1-ylphosphine oxides	SS3
General procedure for the synthesis of diphenyl(2-butyl-3-oxo-2,3-dihydro-2 <i>H</i> -isoindol-1-yl)-phosphine.....	S19
General procedure for the synthesis of diphenyl(2-butyl-3-oxo-2,3-dihydro-2 <i>H</i> -isoindol-1-yl)-phosphine sulphide.....	S19
General procedure for the synthesis of [diphenyl(2-butyl-3-oxo-2,3-dihydro-2 <i>H</i> -isoindol-1-yl)-phosphine] dichloroplatinum	S19
Single crystal X-ray diffraction measurements	S20
¹ H NMR, ¹³ C NMR and ³¹ P NMR spectra	S25
References.....	S69

General information

The reactions were carried out at 25 °C under N₂ atmosphere.

High-performance liquid chromatography-mass spectrometry (HPLC-MS) measurements were performed with an Agilent 1200 liquid chromatography system coupled with a 6130 quadrupole mass spectrometer equipped with an ESI ion source (Agilent Technologies, Palo Alto, CA, USA). Analysis was performed at 40 °C on a Gemini C18 column (150 mm × 4.6 mm, 3 μm; Phenomenex, Torrance, CA, USA) with a mobile phase flow rate of 0.6 mL/min. Composition of eluent A was 0.1% (NH₄)(HCOO) in water; eluent B was 0.1% (NH₄)(HCOO) and 8% water in acetonitrile, 0–3 min 5% B, 3–13 min gradient, 13–20 min 100% B. The injection volume was 2 μL. The chromatographic profile was registered at 254 nm. The MSD operating parameters were as follows: positive ionization mode, scan spectra from m/z 120 to 1000, drying gas temperature 300 °C, nitrogen flow rate 12 L/min, nebulizer pressure 60 psi, capillary voltage 4000 V.

The ³¹P, ¹³C, ¹H NMR spectra were taken in CDCl₃ solution on a Bruker AV-300 or DRX-500 spectrometer operating at 121.5, 75.5 and 300 or 202.4, 125.7 and 500 MHz, respectively. Chemical shifts are downfield relative to 85% H₃PO₄ and TMS. Non-equivalence effects were observed in ¹H and ¹³C{¹H} NMR spectra. Corresponding pairs of resonances were marked with (I) and (II), respectively.

High resolution measurements were performed on a Sciex TripleTOF 5600+ high resolution tandem mass spectrometer equipped with DuoSpray ion source. Electrospray ionization was applied in positive ion detection mode. Samples were dissolved in acetonitrile and flow injected into acetonitrile:water 50:50 flow. The flow rate was 0.2 mL/min. The resolution of the mass spectrometer was 35000.

General procedure for the synthesis of 3-oxoisooindolin-1-ylphosphine oxides (7-20)

A mixture of 1.0 mmol (0.15 g) of 2-formylbenzoic acid, 1.0 mmol of primary amine (0.10 mL of butylamine, 0.12 mL of cyclohexylamine, 0.11 mL of benzylamine or 0.09 mL of aniline) and 1.0 mmol of secondary phosphine oxide (0.20 g of diphenylphosphine oxide, 0.23 g of bis(*p*-tolyl)phosphine oxide, 0.26 g of bis(3,5-dimethylphenyl)phosphine oxide, 0.30 g of bis(2-naphthyl)phosphine oxide, 0.23 g of dibenzylphosphine oxide, 0.18 g of *tert*-butyl(phenyl)phosphine oxide, 0.22 g of 2-methylphenyl(phenyl)phosphine oxide, 0.27 g of 2-trifluoromethylphenyl(phenyl)phosphine oxide, 0.27 g of 3-trifluoromethylphenyl-(phenyl)-phosphine oxide, 0.27 g of 4-trifluoromethylphenyl-(phenyl)phosphine oxide, 0.28 g of 2-phenylphenyl(phenyl)phosphine oxide or 0.25 g of 1-naphthyl(phenyl)phosphine oxide)

was stirred without solvent or in 1 mL of ethanol, toluene or acetonitrile at room temperature under nitrogen atmosphere. The reaction mixtures were analyzed by HPLC chromatography. The solvent and the water formed were eliminated in vacuum, and the crude product so obtained was passed through a 1 cm silica layer using dichloromethane:methanol 97:3 as the eluent. The following products were thus prepared:

Diphenyl (2-butyl-3-oxo-2,3-dihydro-2H-isoindol-1-yl)phosphine oxide (7a)

Yield: 98% (0.38 g), white crystals; Mp: 74–75 °C; ^1H NMR (CDCl_3) δ 0.86 (t, $J_{\text{HH}} = 7.4$, 3H, $\text{CH}_3(\text{CH}_2)_3\text{N}$), 1.10–1.26 (m, 2H, $\text{CH}_2(\text{CH}_2)_2\text{N}$), 1.48–1.66 (m, 2H, $\text{CH}_2\text{CH}_2\text{N}$), 3.17–3.25 (m, 1H, CH_A , CH_2N), 3.92–4.01 (m, 1H, CH_B , CH_2N), 5.49 (d, 1H, $^2J_{\text{HP}} = 11.1$, C_1H), 6.94 (d, 1H, $J_{\text{HH}} = 7.7$, ArH), 7.28–7.36 (m, 1H, ArH), 7.36–7.44 (m, 5H, ArH), 7.44–7.49 (m, 2H, ArH), 7.52–7.60 (m, 2H, ArH), 7.64–7.71 (m, 3H, ArH); ^{13}C NMR (CDCl_3) δ 13.7 ($\text{CH}_3(\text{CH}_2)_3\text{N}$), 19.9 ($\text{CH}_2(\text{CH}_2)_2\text{N}$), 30.0 ($\text{CH}_2\text{CH}_2\text{N}$), 41.7 (CH_2N), 61.0 (d, $^1J_{\text{CP}} = 72.8$, C_1), 123.7 (d, $J_{\text{CP}} = 1.2$, C_4), 124.0 (d, $^3J_{\text{CP}} = 2.4$, C_7), 127.8 (d, $^1J_{\text{CP}} = 81.7$, $\text{C}_1^{''\text{I}}$), 128.6 (d, $^1J_{\text{CP}} = 81.8$, $\text{C}_1^{''\text{II}}$), 128.65 (d, $^3J_{\text{CP}} = 11.7$, $\text{C}_3^{''\text{I}}$), 128.66 (d, $J_{\text{CP}} = 1.9$, C_5), 128.67 (d, $^3J_{\text{CP}} = 11.7$, $\text{C}_3^{''\text{II}}$), 131.1 (d, $J_{\text{CP}} = 2.2$, C_6), 131.6 (d, $^2J_{\text{CP}} = 8.8$, $\text{C}_2^{''\text{I}}$), 131.7 (d, $^2J_{\text{CP}} = 9.0$, $\text{C}_2^{''\text{II}}$), 132.75 (d, $J_{\text{CP}} = 2.9$, $\text{C}_{3\text{a}}$), 132.77 (d, $J_{\text{CP}} = 2.6$, $\text{C}_4^{''\text{I}}$), 132.9 (d, $^3J_{\text{CP}} = 2.8$, $\text{C}_4^{''\text{II}}$), 138.6 (d, $^2J_{\text{CP}} = 2.5$, $\text{C}_{7\text{a}}$), 168.6 (d, $^3J_{\text{CP}} = 1.9$, C_3); ^{31}P (CDCl_3) δ 30.6; $[\text{M}+\text{H}]^+$ found = 390.1614, $\text{C}_{24}\text{H}_{25}\text{NO}_2\text{P}$ requires 390.1622.

Diphenyl (2-cyclohexyl-3-oxo-2,3-dihydro-2H-isoindol-1-yl)phosphine oxide (7b)

Yield: 94% (0.39 g), white crystals; Mp: 59–61 °C; ^1H NMR (CDCl_3) δ 1.13–1.36 (m, 3H, $^c\text{HexH}$), 1.48–1.83 (m, 5H, $^c\text{HexH}$), 2.01–2.22 (m, 1H, $^c\text{HexH}$), 2.36–2.63 (m, 1H, $^c\text{HexH}$), 3.55–3.73 (m, 1H, $\text{C}_1^{''\text{H}}$), 5.42 (d, 1H, $^2J_{\text{HP}} = 11.7$, C_1H), 6.63 (d, 1H, $J_{\text{HH}} = 7.7$, ArH), 7.17–7.43 (m, 7H, ArH), 7.44–7.56 (m, 2H, ArH), 7.56–7.64 (m, 1H, ArH), 7.64–7.82 (m, 3H, ArH); ^{13}C NMR (CDCl_3) δ 25.2 ($\text{C}_4^{''\text{I}}$), 26.0 ($\text{C}_3^{''\text{I}}$), 26.3 ($\text{C}_3^{''\text{II}}$), 29.2 ($\text{C}_2^{''\text{I}}$), 30.0 ($\text{C}_2^{''\text{II}}$), 57.5 ($\text{C}_1^{''\text{I}}$), 63.4 (d, $^1J_{\text{CP}} = 73.4$, C_1), 123.5 (d, $J_{\text{CP}} = 1.1$, C_4), 123.7 (d, $^3J_{\text{CP}} = 2.4$, C_7), 126.8 (d, $^1J_{\text{CP}} = 98.6$, $\text{C}_1^{''\text{II}}$), 128.4 (d, $^3J_{\text{CP}} = 11.8$, $\text{C}_3^{''\text{I}}$), 128.7 (d, $J_{\text{CP}} = 2.6$, C_5), 128.8 (d, $^3J_{\text{CP}} = 11.8$, $\text{C}_3^{''\text{II}}$), 130.1 (d, $^1J_{\text{CP}} = 97.4$, $\text{C}_1^{''\text{II}}$), 130.9 (d, $J_{\text{CP}} = 2.3$, C_6), 131.6 (d, $^2J_{\text{CP}} = 8.8$, $\text{C}_2^{''\text{I}}$), 132.2 (d, $^2J_{\text{CP}} = 8.6$, $\text{C}_2^{''\text{II}}$), 132.7 (d, $J_{\text{CP}} = 2.0$, $\text{C}_4^{''\text{I}}$), 133.8 (d, $J_{\text{CP}} = 2.3$, $\text{C}_4^{''\text{II}}$), 134.4 (d, $^3J_{\text{CP}} = 3.1$, $\text{C}_{3\text{a}}$), 138.9 (d, $^2J_{\text{CP}} = 1.3$, $\text{C}_{7\text{a}}$), 169.0 (d, $^3J_{\text{CP}} = 1.8$, C_3); ^{31}P (CDCl_3) δ 30.8; $[\text{M}+\text{H}]^+$ found = 416.1772, $\text{C}_{26}\text{H}_{27}\text{NO}_2\text{P}$ requires 416.1779.

Diphenyl (2-benzyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (7c)

Yield: 97% (0.41 g), light yellow crystals; Mp: 80–81 °C; ^1H NMR (CDCl_3) δ 4.37 (d, 1H, $J_{\text{HH}} = 15.0$, CH_A , CH_2N), 5.31 (d, 1H, $^2J_{\text{HP}} = 10.7$, C_1H), 5.40 (d, 1H, $J_{\text{HH}} = 15.0$, CH_B , CH_2N), 6.83 (d, 1H, $J_{\text{HH}} = 7.6$, ArH), 7.08–7.20 (m, 2H, ArH), 7.21–7.52 (m, 12H, ArH), 7.52–7.79 (m, 4H, ArH); ^{13}C NMR (CDCl_3) δ 45.3 (CH_2N), 60.2 (d, $^1J_{\text{CP}} = 73.1$, C_1), 124.0 (d, $J_{\text{CP}} = 0.4$, C_4), 124.1 (d, $^3J_{\text{CP}} = 1.8$, C_7), 127.55 (d, $^1J_{\text{CP}} = 97.9$, $\text{C}_1''\text{I}$), 127.59 (C_4'), 128.3 (C_3'), 128.66 (C_2'), 128.69 (d, $^3J_{\text{CP}} = 11.6$, $\text{C}_3''\text{I}$), 128.74 (d, $J_{\text{CP}} = 2.3$, C_5), 128.75 (d, $^3J_{\text{CP}} = 11.7$, $\text{C}_3''\text{II}$), 128.9 (d, $^1J_{\text{CP}} = 97.7$, $\text{C}_1''\text{II}$), 131.3 (d, $J_{\text{CP}} = 2.2$, C_6), 131.6 (d, $^2J_{\text{CP}} = 8.9$, $\text{C}_2''\text{I}$), 132.0 (d, $^2J_{\text{CP}} = 8.9$, $\text{C}_2''\text{II}$), 132.5 (d, $^3J_{\text{CP}} = 3.0$, C_{3a}), 132.8 (d, $J_{\text{CP}} = 2.8$, $\text{C}_4''\text{I}$), 132.9 (d, $J_{\text{CP}} = 2.8$, $\text{C}_4''\text{II}$), 136.6 (C_1'), 138.9 (d, $^2J_{\text{CP}} = 2.4$, C_{7a}), 168.9 (d, $^3J_{\text{CP}} = 1.5$, C_3); ^{31}P (CDCl_3) δ 30.8; $[\text{M}+\text{H}]^+$ found = 424.1457, $\text{C}_{27}\text{H}_{23}\text{NO}_2\text{P}$ requires 424.1466.

Diphenyl (2-phenyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (7d)

Yield: 96% (0.39 g), white crystals; Mp: 268–270 °C; ^1H NMR (CDCl_3) δ 6.50 (t, 1H, $J_{\text{HH}} = 7.2$, ArH), 6.57 (dd, 1H, $J_{\text{HH}} = 5.7$, $^2J_{\text{HP}} = 11.0$, C_1H), 6.83 (d, 2H, $J_{\text{HH}} = 8.0$, ArH), 6.97 (t, 2H, $J_{\text{HH}} = 7.6$, ArH), 7.17–7.28 (m, 4H, ArH), 7.32 (t, 1H, $J_{\text{HH}} = 11.6$, ArH), 7.36–7.42 (m, 1H, ArH), 7.54 (t, 1H, $J_{\text{HH}} = 7.6$, ArH), 7.58–7.67 (m, 4H, ArH), 7.89–7.98 (m, 3H, ArH); ^{13}C NMR (CDCl_3) δ 50.7 (d, $^1J_{\text{CP}} = 76.6$, C_1), 113.9 (d, $J_{\text{CP}} = 0.7$, C_4), 117.6 (d, $^3J_{\text{CP}} = 0.5$, C_7), 127.7 (d, $J_{\text{CP}} = 2.0$, C_5), 128.3 (d, $^3J_{\text{CP}} = 11.6$, $\text{C}_3''\text{I}$), 129.3 (C_4'), 129.4 (d, $^3J_{\text{CP}} = 11.5$, $\text{C}_3''\text{II}$), 129.8 (d, $J_{\text{CP}} = 4.8$, C_3'), 129.9 (d, $J_{\text{CP}} = 5.0$, C_2'), 130.7 (d, $J_{\text{CP}} = 5.5$, C_6), 131.3 (d, $^1J_{\text{CP}} = 97.7$, $\text{C}_1''\text{I}$), 131.36 (d, $^2J_{\text{CP}} = 8.7$, $\text{C}_2''\text{I}$), 131.40 (d, $^2J_{\text{CP}} = 9.0$, $\text{C}_2''\text{II}$), 132.0 (d, $^1J_{\text{CP}} = 96.8$, $\text{C}_1''\text{II}$), 132.1 (d, $^3J_{\text{CP}} = 1.9$, C_{3a}), 132.55 (d, $J_{\text{CP}} = 2.4$, $\text{C}_4''\text{I}$), 132.62 (d, $J_{\text{CP}} = 2.3$, $\text{C}_4''\text{II}$), 139.6 (C_{7a}), 147.6 (d, $^3J_{\text{CP}} = 11.8$, C_1''), 169.1 (d, $^3J_{\text{CP}} = 0.9$, C_3); ^{31}P (CDCl_3) δ 31.8; $[\text{M}+\text{H}]^+$ found = 410.1322, $\text{C}_{26}\text{H}_{21}\text{NO}_2\text{P}$ requires 410.1304.

Bis(*p*-tolyl) (2-butyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (8a)

Yield: 99% (0.41 g), light yellow crystals; Mp: 72–74 °C; ^1H NMR (CDCl_3) δ 0.86 (t, $J_{\text{HH}} = 7.2$, 3H, $\text{CH}_3(\text{CH}_2)_3\text{N}$), 1.10–1.28 (m, 2H, $\text{CH}_2(\text{CH}_2)_2\text{N}$), 1.47–1.69 (m, 2H, $\text{CH}_2\text{CH}_2\text{N}$), 2.37 (s, 3H, $\text{C}_4'\text{CH}_3^{\text{I}}$), 2.41 (s, 3H, $\text{C}_4'\text{CH}_3^{\text{II}}$), 3.20–3.35 (m, 1H, CH_A , CH_2N), 3.89–4.04 (m, 1H, CH_B , CH_2N), 5.44 (d, 1H, $^2J_{\text{HP}} = 11.4$, C_1H), 6.94 (d, 1H, $J_{\text{HH}} = 7.5$, ArH), 7.11–7.44 (m, 9H, ArH), 7.45–7.62 (m, 2H, ArH), 7.68 (d, 1H, $J_{\text{HH}} = 7.3$, ArH); ^{13}C NMR (CDCl_3) δ 13.7 ($\text{CH}_3(\text{CH}_2)_3\text{N}$), 19.9 ($\text{CH}_2(\text{CH}_2)_2\text{N}$), 21.6 ($\text{C}_4'\text{CH}_3^{\text{I}}$), 21.7 ($\text{C}_4'\text{CH}_3^{\text{II}}$), 30.0 ($\text{CH}_2\text{CH}_2\text{N}$), 41.8 (CH_2N), 61.2 (d, $^1J_{\text{CP}} = 72.7$, C_1), 123.6 (d, $J_{\text{CP}} = 1.2$, C_4), 124.1 (d, $^3J_{\text{CP}} = 2.5$, C_7), 124.4 (d,

$^1J_{CP} = 84.5$, C₁’I), 125.8 (d, $^1J_{CP} = 84.3$, C₁’II), 128.5 (d, $J_{CP} = 1.7$, C₅), 129.36 (d, $^3J_{CP} = 12.1$, C₃’I), 129.40 (d, $^3J_{CP} = 12.1$, C₃’II), 131.0 (d, $J_{CP} = 1.8$, C₆), 131.6 (d, $^2J_{CP} = 8.7$, C₂’I), 131.7 (d, $^2J_{CP} = 8.9$, C₂’II), 132.9 (d, $^3J_{CP} = 3.1$, C_{3a}), 138.9 (d, $^2J_{CP} = 2.7$, C_{7a}), 143.4 (d, $J_{CP} = 2.8$, C₄’I), 143.5 (d, $J_{CP} = 2.9$, C₄’II), 168.6 (d, $^3J_{CP} = 1.5$, C₃); ^{31}P (CDCl₃) δ 31.2; [M+H]⁺_{found} = 418.1922, C₂₆H₂₉NO₂P requires 418.1935.

Bis(*p*-tolyl) (2-cyclohexyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (8b)

Yield: 97% (0.43 g), white crystals; Mp: 76–78 °C; 1H NMR (CDCl₃) 1.06–1.16 (m, 1H, ^cHexH), 1.17–1.30 (m, 2H, ^cHexH), 1.53–1.65 (m, 2H, ^cHexH), 1.67–1.80 (m, 3H, ^cHexH), 2.08–2.18 (m, 1H, ^cHexH), 2.34 (s, 3H, C₄”CH₃I), 2.45–2.48 [2.44 (s, C₄”CH₃II), overlapped by the multiplet of ^cHexH total int. 4H], 3.69–3.78 (m, 1H, C₁’H), 5.40 (d, 1H, $^2J_{HP} = 11.9$, C₁H), 6.59 (d, 1H, $J_{HH} = 7.7$, ArH), 7.04–7.13 (m, 4H, ArH), 7.24–7.33 (m, 3H, ArH), 7.36–7.42 (m, 1H, ArH), 7.57–7.64 (m, 2H, ArH), 7.68 (d, 1H, $J_{HH} = 7.7$, ArH); ^{13}C NMR (CDCl₃) δ 21.5 (d, $J_{CP} = 1.2$, C₄”CH₃I), 21.7 (d, $J_{CP} = 1.1$, C₄”CH₃II), 25.2 (C₄’), 26.0 (C₃’I), 26.3 (C₃’II), 29.2 (C₂’I), 30.0 (C₂’II), 57.5 (C₁’), 63.5 (d, $^1J_{CP} = 74.4$, C₁), 123.2 (d, $^1J_{CP} = 100.9$, C₁”I), 123.5 (d, $J_{CP} = 1.2$, C₄’), 123.7 (d, $^3J_{CP} = 2.4$, C₇), 127.3 (d, $^1J_{CP} = 99.8$, C₁”II), 126.6 (d, $J_{CP} = 2.1$, C₅), 129.1 (d, $^3J_{CP} = 12.2$, C₃”I), 129.5 (d, $^3J_{CP} = 12.0$, C₃”II), 130.8 (d, $J_{CP} = 2.4$, C₆), 131.6 (d, $^2J_{CP} = 9.3$, C₂”I), 132.3 (d, $^2J_{CP} = 8.8$, C₂”II), 134.5 (d, $^3J_{CP} = 3.4$, C_{3a}), 139.1 (d, $^2J_{CP} = 1.1$, C_{7a}), 143.30 (C₄”I), 143.33 (C₄”II), 169.0 (d, $^3J_{CP} = 2.0$, C₃’); ^{31}P (CDCl₃) δ 31.7; [M+H]⁺_{found} = 444.2077, C₂₈H₃₁NO₂P requires 444.2086.

Bis(*p*-tolyl) (2-benzyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (8c)

Yield: 97% (0.44 g), light yellow crystals; Mp: 80–82 °C; 1H NMR (CDCl₃) δ 2.39 (s, 3H, C₄”CH₃I), 2.41 (s, 3H, C₄”CH₃II), 4.45 (d, 1H, $J_{HH} = 14.9$, CH_A, CH₂N), 5.27 (d, 1H, $^2J_{HP} = 10.9$, C₁H), 5.38 (d, 1H, $J_{HH} = 14.9$, CH_B, CH₂N), 6.82 (d, 1H, $J_{HH} = 7.7$, ArH), 7.08–7.45 (m, 13H, ArH), 7.46–7.64 (m, 2H, ArH), 7.74 (d, 1H, $J_{HH} = 7.5$, ArH); ^{13}C NMR (CDCl₃) δ 21.6 (C₄”CH₃I), 21.7 (C₄”CH₃II), 45.2 (CH₂N), 60.3 (d, $^1J_{CP} = 73.1$, C₁), 124.0 (d, $J_{CP} = 1.0$, C₄’), 124.06 (d, $^3J_{CP} = 2.3$, C₇), 124.12 (d, $^1J_{CP} = 101.1$, C₁”I), 126.0 (d, $^1J_{CP} = 100.1$, C₁”II), 127.5 (C₄’), 128.4 (C₃’), 128.60 (d, $J_{CP} = 3.3$, C₅), 128.61 (C₂’), 129.36 (d, $^3J_{CP} = 12.0$, C₃”I), 129.44 (d, $^3J_{CP} = 12.2$, C₃”II), 131.2 (d, $J_{CP} = 1.8$, C₆), 131.6 (d, $^2J_{CP} = 9.3$, C₂”I), 132.0 (d, $^2J_{CP} = 9.2$, C₂”II), 132.6 (d, $^3J_{CP} = 2.9$, C_{3a}), 136.8 (C₁’), 139.1 (d, $^2J_{CP} = 2.1$, C_{7a}), 143.4 (d, $J_{CP} = 2.8$,

$C_4^{''I}$), 143.7 (d, $J_{CP} = 3.0$, $C_4^{''II}$), 168.8 (d, $^3J_{CP} = 1.6$, C_3); ^{31}P ($CDCl_3$) δ 31.4; $[M+H]^+$ found = 452.1765, $C_{29}H_{27}NO_2P$ requires 452.1779.

Bis(*p*-tolyl) (2-phenyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (8d)

Yield: 98% (0.43 g), white crystals; Mp: 188–190 °C; 1H NMR ($CDCl_3$) δ 2.22 (s, 3H, $C_4^{''}CH_3^I$), 2.36 (s, 3H, $C_4^{''}CH_3^{II}$), 6.41–6.55 [6.50 (d, 2H, $^2J_{HP} = 6.8$, C_1H) overlapped by the multiplet of ArH, total int. 2H], 6.80 (d, 2H, $J_{HH} = 8.1$, ArH), 6.97 (t, 2H, $J_{HH} = 7.7$, ArH), 7.00–7.15 (m, 3H, ArH), 7.17–7.31 (m, 2H, ArH), 7.40 (dd, 2H, $J_{HH} = 8.1$, $J_{HH} = 2.8$, ArH), 7.52 (t, 1H, $J_{HH} = 7.4$, ArH), 7.67 (d, 1H, $J_{HH} = 7.8$, ArH), 7.78 (d, 1H, $J_{HH} = 7.8$, ArH), 7.81 (d, 1H, $J_{HH} = 7.9$, ArH), 7.89 (d, 1H, $J_{HH} = 7.9$, ArH); ^{13}C NMR ($CDCl_3$) δ 21.5 ($C_4^{''}CH_3^I$), 21.6 ($C_4^{''}CH_3^{II}$), 50.8 (d, $^1J_{CP} = 77.0$, C_1), 113.8 (d, $J_{CP} = 0.6$, C_4), 117.5 (d, $^3J_{CP} = 0.5$, C_7), 127.6 (d, $J_{CP} = 1.7$, C_5), 128.4 (d, $^1J_{CP} = 101.3$, $C_1^{''I}$), 128.9 (d, $^3J_{CP} = 11.8$, $C_3^{''I}$), 129.0 (d, $^1J_{CP} = 99.0$, $C_1^{''II}$), 129.3 (C_4'), 129.8 (d, $J_{CP} = 4.3$, C_3'), 129.9 (d, $^3J_{CP} = 12.0$, $C_3^{''II}$), 130.1 (d, $J_{CP} = 5.0$, C_2'), 130.7 (d, $J_{CP} = 2.2$, C_6), 131.4 (d, $^2J_{CP} = 9.0$, $C_2^{''I}$), 131.5 (d, $^2J_{CP} = 9.5$, $C_2^{''II}$), 132.4 (d, $J_{CP} = 2.2$, C_{3a}), 139.8 (C_{7a}), 142.0 (d, $J_{CP} = 2.8$, $C_4^{''I}$), 142.6 (d, $J_{CP} = 2.7$, $C_4^{''II}$), 147.7 (d, $^3J_{CP} = 12.0$, C_1'), 169.1 (d, $^3J_{CP} = 1.5$, C_3); ^{31}P ($CDCl_3$) δ 31.3; $[M+H]^+$ found = 438.1637, $C_{28}H_{25}NO_2P$ requires 438.1617.

Bis(3,5-dimethylphenyl) (2-butyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (9a)

Yield: 97% (0.43 g), yellow crystals; Mp: 168–170 °C; 1H NMR ($CDCl_3$) δ 0.89 (t, $J_{HH} = 7.3$, 3H, $CH_3(CH_2)_3N$), 1.14–1.30 (m, 2H, $CH_2(CH_2)_2N$), 1.47–1.72 (m, 2H, CH_2CH_2N), 2.23 (s, 6H, $C_3'CH_3^I$), 2.34 (s, 6H, $C_3'CH_3^{II}$), 3.25–3.37 (m, 1H, CH_A , CH_2N), 3.90–4.04 (m, 1H, CH_B , CH_2N), 5.47 (d, 1H, $^2J_{HP} = 10.9$, C_1H), 6.83–6.94 (m, 2H, ArH) 7.12–7.45, (m, 7H, ArH), 7.71 (d, 1H, $J_{HH} = 7.4$, ArH); ^{13}C NMR ($CDCl_3$) δ 13.8 ($CH_3(CH_2)_3N$), 20.0 ($CH_2(CH_2)_2N$), 21.2 ($C_3'CH_3^I$), 21.3 ($C_3'CH_3^{II}$), 30.2 (CH_2CH_2N), 41.8 (CH_2N), 61.2 (d, $^1J_{CP} = 71.8$, C_1), 123.5 (d, $J_{CP} = 1.2$, C_4), 124.2 (d, $^3J_{CP} = 2.3$, C_7), 127.2 (d, $^1J_{CP} = 97.5$, $C_1^{''I}$), 128.6 (d, $J_{CP} = 1.6$, C_5), 129.1 (d, $^1J_{CP} = 97.0$, $C_1^{''II}$), 129.2 (d, $^2J_{CP} = 9.0$, $C_2^{''I}$), 129.4 (d, $^2J_{CP} = 8.9$, $C_2^{''II}$), 130.9 (d, $J_{CP} = 2.2$, C_6), 133.2 (d, $^3J_{CP} = 3.1$, C_{3a}), 134.46 (d, $J_{CP} = 3.1$, $C_4^{''I}$), 134.54 (d, $J_{CP} = 2.9$, $C_4^{''II}$), 138.3 (d, $^3J_{CP} = 12.4$, $C_3^{''I}$), 138.5 (d, $^3J_{CP} = 12.3$, $C_3^{''II}$), 138.9 (d, $^2J_{CP} = 2.3$, C_{7a}), 168.6 (d, $^3J_{CP} = 1.5$, C_3); ^{31}P ($CDCl_3$) δ 31.6; $[M+H]^+$ found = 446.2237, $C_{28}H_{33}NO_2P$ requires 446.2248.

Bis(3,5-dimethylphenyl) (2-cyclohexyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (9b)

Yield: 96% (0.45 g), white crystals; Mp: 180–182 °C; ^1H NMR (CDCl_3) δ 1.15–1.32 (m, 3H, $^c\text{HexH}$), 1.51–1.67 (m, 2H, $^c\text{HexH}$), 1.67–1.83 (m, 3H, $^c\text{HexH}$), 2.04–2.22 [2.18 (s, $\text{C}_3''\text{CH}_3^{\text{I}}$), overlapped by the multiplet of $^c\text{HexH}$ total int. 7H], 2.35 (s, 6H, $\text{C}_3''\text{CH}_3^{\text{II}}$), 2.38–2.51 (m, 1H, $^c\text{HexH}$), 3.62–3.76 (m, 1H, $\text{C}_1'\text{H}$), 5.40 (d, 1H, $^2J_{\text{HP}} = 11.3$, C_1H), 6.60 (d, 1H, $J_{\text{HH}} = 7.7$, ArH), 6.77 (d, 2H, $J_{\text{HH}} = 11.3$, ArH), 7.11 (s, 1H, ArH), 7.19–7.44 (m, 5H, ArH), 7.69 (d, 1H, $J_{\text{HH}} = 7.7$, ArH); ^{13}C NMR (CDCl_3) δ 21.1 ($\text{C}_3''\text{CH}_3^{\text{I}}$), 21.3 ($\text{C}_3''\text{CH}_3^{\text{II}}$), 25.2 (C_4'), 26.0 ($\text{C}_3''\text{I}$), 26.3 ($\text{C}_3''\text{II}$), 29.2 ($\text{C}_2''\text{I}$), 29.9 ($\text{C}_2''\text{II}$), 57.4 (C_1'), 63.4 (d, $^1J_{\text{CP}} = 72.4$, C_1), 123.3 (d, $J_{\text{CP}} = 1.4$, C_4), 123.8 (d, $^3J_{\text{CP}} = 2.2$, C_7), 126.0 (d, $^1J_{\text{CP}} = 98.0$, $\text{C}_1''\text{I}$), 128.5 (d, $J_{\text{CP}} = 2.0$, C_5), 129.1 (d, $^2J_{\text{CP}} = 9.0$, $\text{C}_2''\text{I}$), 129.9 (d, $^2J_{\text{CP}} = 8.5$, $\text{C}_2''\text{II}$), 130.4 (d, $^1J_{\text{CP}} = 96.3$, $\text{C}_1''\text{II}$), 130.7 (d, $J_{\text{CP}} = 2.3$, C_6), 134.32 (d, $J_{\text{CP}} = 3.0$, $\text{C}_4''\text{I}$), 134.34 (d, $J_{\text{CP}} = 2.9$, $\text{C}_4''\text{II}$), 134.6 (d, $^3J_{\text{CP}} = 3.1$, $\text{C}_{3\text{a}}$), 137.8 (d, $^3J_{\text{CP}} = 12.4$, $\text{C}_3''\text{I}$), 138.5 (d, $^3J_{\text{CP}} = 12.3$, $\text{C}_3''\text{II}$), 139.1 (d, $^2J_{\text{CP}} = 1.4$, $\text{C}_{7\text{a}}$), 168.9 (d, $^3J_{\text{CP}} = 2.1$, C_3); ^{31}P (CDCl_3) δ 31.7; $[\text{M}+\text{H}]^+_{\text{found}} = 472.2378$, $\text{C}_{30}\text{H}_{35}\text{NO}_2\text{P}$ requires 472.2399.

Bis(3,5-dimethylphenyl) (2-benzyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (9c)

Yield: 97% (0.46 g), white crystals; Mp: 88–90 °C; ^1H NMR (CDCl_3) δ 2.23 (s, 6H, $\text{C}_3''\text{CH}_3^{\text{I}}$), 2.31 (s, 6H, $\text{C}_3''\text{CH}_3^{\text{II}}$), 4.35 (d, 1H, $J_{\text{HH}} = 15.1$, CH_A , CH_2N), 5.29 (d, 1H, $^2J_{\text{HP}} = 10.0$, C_1H), 5.39 (d, 1H, $J_{\text{HH}} = 15.1$, CH_B , CH_2N), 6.83 (d, 1H, $J_{\text{HH}} = 7.7$, ArH), 6.91 (d, 2H, $J_{\text{HH}} = 11.5$, ArH), 7.07–7.46 (m, 11H, ArH), 7.77 (d, 1H, $J_{\text{HH}} = 7.5$, ArH); ^{13}C NMR (CDCl_3) δ 21.2 ($\text{C}_3''\text{CH}_3^{\text{I}}$), 21.3 ($\text{C}_3''\text{CH}_3^{\text{II}}$), 45.2 (CH_2N), 60.1 (d, $^1J_{\text{CP}} = 71.7$, C_1), 123.8 (d, $J_{\text{CP}} = 1.2$, C_4), 124.2 (d, $^3J_{\text{CP}} = 2.1$, C_7), 127.39 (d, $^1J_{\text{CP}} = 97.6$, $\text{C}_1''\text{I}$), 127.44 (C_4'), 128.2 (C_3'), 128.56 (d, $J_{\text{CP}} = 2.0$, C_5), 128.58 (C_2'), 129.0 (d, $^1J_{\text{CP}} = 96.9$, $\text{C}_1''\text{II}$), 129.1 (d, $^2J_{\text{CP}} = 8.9$, $\text{C}_2''\text{I}$), 129.5 (d, $^2J_{\text{CP}} = 8.9$, $\text{C}_2''\text{II}$), 131.1 (d, $J_{\text{CP}} = 1.8$, C_6), 132.7 (d, $^3J_{\text{CP}} = 2.8$, $\text{C}_{3\text{a}}$), 134.4 (d, $J_{\text{CP}} = 3.0$, $\text{C}_4''\text{I}$), 134.5 (d, $J_{\text{CP}} = 2.9$, $\text{C}_4''\text{II}$), 136.8 (C_1'), 138.3 (d, $^3J_{\text{CP}} = 12.5$, $\text{C}_3''\text{I}$), 138.5 (d, $^3J_{\text{CP}} = 12.4$, $\text{C}_3''\text{II}$), 139.1 (d, $^2J_{\text{CP}} = 2.2$, $\text{C}_{7\text{a}}$), 168.8 (d, $^3J_{\text{CP}} = 1.5$, C_3); ^{31}P (CDCl_3) δ 31.3; $[\text{M}+\text{H}]^+_{\text{found}} = 480.2086$, $\text{C}_{31}\text{H}_{31}\text{NO}_2\text{P}$ requires 480.2092.

Bis(3,5-dimethylphenyl) (2-phenyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (9d)

Yield: 97% (0.45 g), white crystals; Mp: 200–202 °C; ^1H NMR (CDCl_3) δ 2.09 (s, 6H, $\text{C}_3''\text{CH}_3^{\text{I}}$), 2.31 (s, 6H, $\text{C}_3''\text{CH}_3^{\text{II}}$), 6.41 (dd, 1H, $^2J_{\text{HP}} = 10.6$, $J_{\text{HH}} = 6.4$, C_1H), 6.49 (t, 1H, $J_{\text{HH}} = 7.1$, ArH), 6.81 (t, 3H, $J_{\text{HH}} = 9.8$, ArH), 6.92–7.03 (m, 3H, ArH), 7.18–7.31 (m, 3H, ArH), 7.46–7.59 (m, 3H, ArH), 7.65 (d, 1H, $J_{\text{HH}} = 7.8$, ArH), 7.87 (d, 1H, $J_{\text{HH}} = 7.9$, ArH);

¹³C NMR (CDCl₃) δ 21.2 (C₃''CH₃^I), 21.5 (C₃''CH₃^{II}), 50.8 (d, ¹J_{CP} = 75.6, C₁), 113.9 (d, J_{CP} = 2.6, C₄), 117.5 (d, ³J_{CP} = 1.2, C₇), 127.5 (d, J_{CP} = 2.5, C₅), 128.90 (d, ²J_{CP} = 9.0, C₂^{''I}), 128.92 (d, ²J_{CP} = 9.0, C₂^{''II}), 129.3 (C₄'), 129.6 (d, J_{CP} = 4.8, C₃'), 129.9 (d, J_{CP} = 4.9, C₂'), 130.7 (d, J_{CP} = 1.3, C₆), 131.1 (d, ¹J_{CP} = 97.3, C₁^{''I}), 131.9 (d, ¹J_{CP} = 95.3, C₁^{''II}), 132.5 (d, ³J_{CP} = 2.4, C_{3a}), 133.4 (d, J_{CP} = 2.6, C₄^{''I}), 134.0 (d, J_{CP} = 2.9, C₄^{''II}), 137.4 (d, ³J_{CP} = 12.1, C₃^{''I}), 138.4 (d, ³J_{CP} = 12.0, C₃^{''II}), 140.1 (C_{7a}), 147.6 (d, ³J_{CP} = 11.5, C₁'), 169.3 (d, ³J_{CP} = 1.1, C₃); ³¹P (CDCl₃) δ 32.2; [M+H]⁺_{found} = 466.1949, C₃₀H₂₉NO₂P requires 466.1930.

Bis(2-naphthyl) (2-butyl-3-oxo-2,3-dihydro-2H-isoindol-1-yl)phosphine oxide (10a)

Yield: 96% (0.47 g), light yellow crystals; Mp: 105–107 °C; ¹H NMR (CDCl₃) δ 0.81 (t, 3H, J_{HH} = 7.4, CH₃(CH₂)₃N), 1.07–1.23 (m, 2H, CH₂(CH₂)₂N), 1.52–1.71 (m, 2H, CH₂CH₂N), 3.30–3.39 (m, 1H, CH_A, CH₂N), 3.94–4.03 (m, 1H, CH_B, CH₂N), 5.68 (d, 1H, ²J_{HP} = 11.1, C₁H), 6.95 (d, 1H, J_{HH} = 7.7, ArH), 7.20 (t, 1H, J_{HH} = 8.8, ArH), 7.25–7.31 (m, 1H, ArH), 7.37 (t, 1H, J_{HH} = 7.5, ArH), 7.52–7.66 (m, 4H, ArH), 7.67 (d, 1H, J_{HH} = 7.5, ArH), 7.72 (t, 2H, J_{HH} = 8.7, ArH), 7.76–7.83 (m, 2H, ArH), 7.86 (d, 2H, J_{HH} = 8.2, ArH), 7.90 (d, 1H, J_{HH} = 8.2, ArH), 7.95 (dd, 1H, J_{HH} = 8.5, J_{HH} = 3.1, ArH), 8.15 (d, 1H, J_{HH} = 13.5, ArH), 8.29 (d, 1H, J_{HH} = 13.4, ArH); ¹³C NMR (CDCl₃) δ 13.6 (CH₃(CH₂)₃N), 19.9 (CH₂(CH₂)₂N), 30.1 (CH₂CH₂N), 41.9 (CH₂N), 61.2 (d, ¹J_{CP} = 73.0, C₁), 123.8 (d, J_{CP} = 1.2, C₄), 124.1 (d, ³J_{CP} = 2.3, C₇), 124.9 (d, ¹J_{CP} = 98.4, C₂^{''I}), 125.7 (d, J_{CP} = 9.9, C^{''I}), 125.9 (d, J_{CP} = 10.0, C^{''II}), 126.3 (d, ¹J_{CP} = 98.1, C₂^{''II}), 127.2 (C^{''I}), 127.3 (C^{''II}), 127.8 (C^{''I}), 127.9 (C^{''II}), 128.4 (d, J_{CP} = 11.5, C^{''I}), 128.6 (d, J_{CP} = 11.5, C^{''II}), 128.7 (C^{''I}), 128.75 (d, J_{CP} = 1.0, C₅), 128.77 (C^{''II}), 128.9 (C^{''I}), 129.0 (C^{''II}), 131.2 (d, J_{CP} = 2.3, C₆), 132.35 (d, ²J_{CP} = 12.9, C^{''I}), 132.39 (d, ²J_{CP} = 13.0, C^{''II}), 132.9 (d, ³J_{CP} = 3.1, C_{3a}), 134.2 (d, ²J_{CP} = 8.5, C^{''I}), 134.7 (d, ²J_{CP} = 8.5, C^{''II}), 134.95 (d, J_{CP} = 2.3, C^{''I}), 135.02 (d, J_{CP} = 2.5, C^{''II}), 138.7 (d, ²J_{CP} = 2.4, C_{7a}), 168.6 (d, ³J_{CP} = 1.6, C₃); ³¹P (CDCl₃) δ 31.2; [M+H]⁺_{found} = 490.1930, C₃₂H₂₉NO₂P requires 490.1930.

Bis(2-naphthyl) (2-cyclohexyl-3-oxo-2,3-dihydro-2H-isoindol-1-yl)phosphine oxide (10b)

Yield: 95% (0.49 g), light yellow crystals; Mp: 208–210 °C; ¹H NMR (CDCl₃) δ 0.79–1.08 (m, 1H, ^cHexH), 1.13–1.37 (m, 2H, ^cHexH), 1.48–1.61 (m, 1H, ^cHexH), 1.61–1.90 (m, 4H, ^cHexH), 2.09–2.26 (m, 1H, ^cHexH), 2.38–2.57 (m, 1H, ^cHexH), 3.71–3.89 (m, 1H, C₁'H),

5.64 (d, 1H, $^2J_{\text{HP}} = 11.5$, C₁H), 6.66 (d, 1H, $J_{\text{HH}} = 7.7$, ArH), 7.10 (t, 1H, $J_{\text{HH}} = 8.8$, ArH), 7.19–7.32 (m, 1H, ArH), 7.41 (t, 1H, $J_{\text{HH}} = 7.6$, ArH), 7.51–7.73 (m, 5H, ArH), 7.73–7.82 (m, 2H, ArH), 7.82–7.92 (m, 2H, ArH), 7.95 (d, 1H, $J_{\text{HH}} = 8.2$, ArH), 7.97–8.07 (m, 2H, ArH), 8.32 (d, 1H, $J_{\text{HH}} = 13.3$, ArH); ^{13}C NMR (CDCl₃) δ 25.2 (C_{4'}), 26.0 (C_{3''I}), 26.2 (C_{3''II}), 29.3 (C_{2''I}), 30.1 (C_{2''II}), 57.7 (C_{1'}), 63.5 (d, $^1J_{\text{CP}} = 73.5$, C₁), 123.6 (d, $^1J_{\text{CP}} = 92.0$, C_{1''I}), 123.7 (d, $J_{\text{CP}} = 1.8$, C₄), 123.8 (d, $^3J_{\text{CP}} = 2.4$, C₇), 126.1 (d, $J_{\text{CP}} = 9.8$, C_{''I}), 126.3 (d, $J_{\text{CP}} = 9.4$, C_{''II}), 127.1 (C_{''I}), 127.4 (C_{''II}), 127.7 (d, $J_{\text{CP}} = 97.6$, C_{2''I}), 128.1 (d, $J_{\text{CP}} = 11.6$, C_{''I}), 128.76 (d, $J_{\text{CP}} = 11.2$, C_{''II}), 128.78 (C_{''I}), 128.84 (d, $J_{\text{CP}} = 1.8$, C₅), 128.96 (C_{''II}), 128.99 (C_{''I}), 129.1 (C_{''II}), 131.1 (d, $J_{\text{CP}} = 2.3$, C₆), 132.3 (d, $J_{\text{CP}} = 12.9$, C_{''I}), 132.5 (d, $J_{\text{CP}} = 12.8$, C_{''II}), 134.0 (d, $J_{\text{CP}} = 8.8$, C_{''I}), 134.6 (d, $^3J_{\text{CP}} = 3.0$, C_{3a}), 135.02 (d, $J_{\text{CP}} = 2.1$, C_{''I}), 135.04 (d, $J_{\text{CP}} = 2.2$, C_{''II}), 135.3 (d, $J_{\text{CP}} = 7.8$, C_{''II}), 138.9 (d, $^2J_{\text{CP}} = 1.3$, C_{7a}), 169.0 (d, $^3J_{\text{CP}} = 2.0$, C₃); ^{31}P (CDCl₃) δ 31.4; [M+H]⁺_{found} = 516.2104, C₃₄H₃₁NO₂P requires 516.2086.

Bis(2-naphthyl) (2-benzyl-3-oxo-2,3-dihydro-2H-isoindol-1-yl)phosphine oxide (10c)

Yield: 97% (0.51 g), light yellow crystals; Mp: 174–176 °C; ^1H NMR (CDCl₃) δ 4.48 (d, 1H, $J_{\text{HH}} = 15.0$, CH_A, CH₂N), 5.44 (d, 1H, $J_{\text{HH}} = 15.0$, CH_B, CH₂N), 5.53 (d, 1H, $^2J_{\text{HP}} = 10.4$, C₁H), 6.90 (d, 1H, $J_{\text{HH}} = 7.7$, ArH), 7.11–7.22 (m, 3H, ArH), 7.23–7.34 (m, 4H, ArH), 7.41 (t, 2H, $J_{\text{HH}} = 7.5$, ArH), 7.53–8.01 (m, 12H, ArH), 8.20 (d, 1H, $J_{\text{HH}} = 13.4$, ArH), 8.27 (d, 1H, $J_{\text{HH}} = 13.4$, ArH); ^{13}C NMR (CDCl₃) δ 45.5 (CH₂N), 60.4 (d, $^1J_{\text{CP}} = 73.4$, C₁), 124.25 (d, $^3J_{\text{CP}} = 1.6$, C₇), 124.28 (d, $J_{\text{CP}} = 0.5$, C₄), 125.0 (d, $^1J_{\text{CP}} = 98.6$, C_{2''I}), 125.96 (d, $J_{\text{CP}} = 9.9$, C_{''I}), 126.00 (d, $J_{\text{CP}} = 9.6$, C_{''II}), 127.27 (C_{''I}), 127.33 (C_{''II}), 127.7 (C_{4'}), 127.8 (d, $^1J_{\text{CP}} = 92.1$, C_{2''II}), 127.9 (C_{''I}), 128.0 (C_{''II}), 128.40 (C_{2'}), 128.42 (d, $J_{\text{CP}} = 11.8$, C_{''I}), 128.71 (d, $J_{\text{CP}} = 11.8$, C_{''II}), 128.74 (C_{3'}), 128.78 (C_{''I}), 128.80 (C_{''II}), 128.86 (C_{''I}), 128.88 (C_{''II}), 129.1 (d, $J_{\text{CP}} = 1.0$, C₅), 131.5 (d, $J_{\text{CP}} = 2.0$, C₆), 132.4 (d, $J_{\text{CP}} = 12.9$, C_{''I}), 132.5 (d, $J_{\text{CP}} = 13.1$, C_{''II}), 132.6 (d, $^3J_{\text{CP}} = 3.0$, C_{3a}), 134.2 (d, $J_{\text{CP}} = 8.6$, C_{''I}), 135.0 (d, $J_{\text{CP}} = 2.5$, C_{''I}), 135.06 (d, $J_{\text{CP}} = 8.3$, C_{''II}), 135.13 (d, $J_{\text{CP}} = 2.5$, C_{''II}), 136.7 (C_{1'}), 139.0 (d, $^2J_{\text{CP}} = 2.3$, C_{7a}), 167.0 (d, $^3J_{\text{CP}} = 2.0$, C₃); ^{31}P (CDCl₃) δ 31.1; [M+H]⁺_{found} = 524.1793, C₃₅H₂₇NO₂P requires 524.1773.

Dibenzyl (2-butyl-3-oxo-2,3-dihydro-2H-isoindol-1-yl)phosphine oxide (11a)

Yield: 96% (0.40 g), colourless oil; ^1H NMR (CDCl₃) δ 0.85 (t, $J_{\text{HH}} = 7.3$, 3H, CH₃(CH₂)₃N), 1.13–1.30 (m, 2H, CH₂(CH₂)₂N), 1.36–1.55 (m, 2H, CH₂CH₂N), 2.71–2.95 (m, 2H, CH₂P^I), 3.00–3.19 (m, 2H, CH₂P^{II}), 3.47–3.59 (m, 1H, CH_A, CH₂N), 3.98–4.11 (m, 1H, CH_B, CH₂N),

4.96 (d, 1H, $^2J_{HP} = 8.9$, C₁H), 7.03–7.13 (m, 2H, ArH), 7.17–7.32 (m, 8H, ArH), 7.49–7.64 (m, 2H, ArH), 7.71 (d, 1H, $J_{HH} = 7.4$, ArH), 7.89 (d, 1H, $J_{HH} = 7.2$, ArH); ^{13}C NMR (CDCl₃) δ 13.7 (CH₃(CH₂)₃N), 19.9 (CH₂(CH₂)₂N), 30.0 (CH₂CH₂N), 33.5 (d, $^1J_{CP} = 60.1$, CH₂P^I), 33.8 (d, $^1J_{CP} = 59.0$, CH₂P^{II}), 42.3 (CH₂N), 58.4 (d, $^1J_{CP} = 65.0$, C₁), 123.8 (d, $J_{CP} = 2.2$, C₄), 124.4 (d, $^3J_{CP} = 1.3$, C₇), 127.17 (d, $J_{CP} = 2.7$, C_{4'}^I), 127.24 (d, $J_{CP} = 2.5$, C_{4'}^{II}), 128.76 (d, $J_{CP} = 12.1$, C_{3'}^I), 128.79 (d, $J_{CP} = 12.1$, C_{3'}^{II}), 129.0 (d, $J_{CP} = 1.9$, C₅), 129.7 (d, $^3J_{CP} = 7.7$, C_{2'}^I), 129.8 (d, $^3J_{CP} = 7.6$, C_{2'}^{II}), 130.1 (d, $^2J_{CP} = 11.4$, C_{1'}^I), 130.2 (d, $^2J_{CP} = 11.4$, C_{1'}^{II}), 131.7 (d, $J_{CP} = 2.1$, C₆), 132.7 (d, $^3J_{CP} = 2.8$, C_{3a}), 138.8 (d, $^2J_{CP} = 2.0$, C_{7a}), 168.7 (d, $^3J_{CP} = 2.0$, C₃); ^{31}P (CDCl₃) δ 41.7; [M+H]⁺_{found} = 418.1942, C₂₆H₂₉NO₂P requires 418.1936.

Dibenzyl (2-cyclohexyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (11b)

Yield: 94% (0.42 g), white crystals; Mp: 64–65 °C; 1H NMR (CDCl₃) δ 1.16–1.45 (m, 3H, ^cHexH), 1.52–1.69 (m, 2H, ^cHexH), 1.70–2.07 (m, 3H, ^cHexH), 2.11–2.30, (m, 1H, ^cHexH), 2.46–2.72 (m, 2H, ^cHexH, CH_A, CH₂P^I), 2.81–3.00 (m, 1H, CH_B, CH₂P^I), 3.02–3.24 (m, 2H, CH₂P^{II}), 3.60–3.76 (m, C_{1'}H), 4.94 (d, 1H, $^2J_{HP} = 8.9$, C₁H), 6.97–7.09 (m, 2H, ArH), 7.17–7.32 (m, 8H, ArH), 7.47–7.63 (m, 2H, ArH), 7.68 (d, 1H, $J_{HH} = 7.5$, ArH), 7.87 (d, 1H, $J_{HH} = 7.0$, ArH); ^{13}C NMR (CDCl₃) δ 25.1 (C_{4'}), 25.9 (C_{3'}^I), 26.3 (C_{3'}^{II}), 28.8 (C_{2'}^I), 30.0 (C_{2'}^{II}), 32.7 (d, $^1J_{CP} = 60.6$, CH₂P^I), 33.3 (d, $^1J_{CP} = 59.3$, CH₂P^{II}), 57.9 (C_{1'}), 60.9 (d, $^1J_{CP} = 64.7$, C₁), 123.5 (d, $J_{CP} = 2.2$, C₄), 124.2 (d, $J_{CP} = 1.2$, C₇), 127.1 (d, $J_{CP} = 2.7$, C_{4'}^I), 127.2 (d, $J_{CP} = 2.5$, C_{4'}^{II}), 128.75 (d, $J_{CP} = 13.6$, C_{3'}^I), 128.78 (d, $J_{CP} = 13.5$, C_{3'}^{II}), 129.0 (d, $J_{CP} = 1.9$, C₅), 129.83 (d, $^3J_{CP} = 5.5$, C_{2'}^I), 129.84 (d, $^3J_{CP} = 5.6$, C_{2'}^{II}), 130.2 (d, $^2J_{CP} = 10.7$, C_{1'}^I), 130.3 (d, $^2J_{CP} = 10.7$, C_{1'}^{II}), 131.6 (d, $J_{CP} = 2.2$, C₆), 134.1 (d, $^3J_{CP} = 3.0$, C_{3a}), 139.1 (d, $^2J_{CP} = 1.4$, C_{7a}), 169.2 (d, $^3J_{CP} = 2.2$, C₃); ^{31}P (CDCl₃) δ 41.4; [M+H]⁺_{found} = 444.2098, C₂₈H₃₁NO₂P requires 444.2092.

Dibenzyl (2-benzyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (11c)

Yield: 96% (0.43 g), white crystals; Mp: 68–70 °C; 1H NMR (CDCl₃) δ 2.54–2.68 (m, 1H, CH_A, CH₂P^I), 2.77–2.91 (m, 1H, CH_B, CH₂P^I), 3.07–3.23 (m, 2H, CH₂P^{II}), 4.77 (d, 1H, $J_{HH} = 14.7$, CH_A, CH₂N), 4.79 (d, 1H, $^2J_{HP} = 8.8$, C₁H), 5.34 (d, 1H, $J_{HH} = 14.6$, CH_B, CH₂N), 6.95–7.04 (m, 2H, ArH), 7.13–7.31 (m, 13H, ArH), 7.51–7.60 (m, 3H, ArH), 7.94 (d, 1H, $J_{HH} = 6.3$, ArH); ^{13}C NMR (CDCl₃) δ 33.0 (d, $^1J_{CP} = 60.3$, CH₂P^I), 34.4 (d, $^1J_{CP} = 58.4$, CH₂P^{II}), 45.9 (CH₂N), 57.5 (d, $^1J_{CP} = 65.3$, C₁), 123.2 (d, $J_{CP} = 2.1$, C₄), 124.9 (d, $J_{CP} = 1.1$,

C_7), 127.1 (d, $J_{CP} = 2.5$, $C_4''^I$), 127.3 (d, $J_{CP} = 2.5$, $C_4''^{\text{II}}$), 127.5 (C_4'), 128.6 (C_3'), 128.75 (d, $J_{CP} = 16.0$, $C_3''^I$), 128.77 (d, $J_{CP} = 15.8$, $C_3''^{\text{II}}$), 128.8 (C_2'), 129.0 (d, $J_{CP} = 1.9$, C_5), 129.71 (d, $^3J_{CP} = 5.5$, $C_2''^I$), 129.8 (d, $^3J_{CP} = 5.4$, $C_2''^{\text{II}}$), 129.9 (d, $^2J_{CP} = 17.8$, $C_1''^I$), 130.0 (d, $^2J_{CP} = 17.6$, $C_1''^{\text{II}}$), 131.9 (d, $J_{CP} = 2.2$, C_6), 132.5 (d, $^3J_{CP} = 2.8$, C_{3a}), 136.6 (C_1'), 139.0 (d, $^2J_{CP} = 2.0$, C_{7a}), 168.6 (d, $^3J_{CP} = 2.2$, C_3); ^{31}P (CDCl_3) δ 41.5; $[\text{M}+\text{H}]^+_{\text{found}} = 452.1783$, $\text{C}_{29}\text{H}_{27}\text{NO}_2\text{P}$ requires 452.1779.

***tert*-Butyl(phenyl) (2-butyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (12)**

Yield: 94% (0.44 g), *dr*: 40:60, colourless oil; characterized as a mixture, **Isomer A:** ^1H NMR (CDCl_3) δ 0.78 (t, $J_{HH} = 7.3$, 3H, $\text{CH}_3(\text{CH}_2)_3\text{N}$), 1.00–1.27 [1.22 (d, $^3J_{HP} = 14.9$, $\text{C}(\text{CH}_3)_3$) overlapped by the multiplet of $\text{CH}_2(\text{CH}_2)_2\text{N}$, total int. 11H], 1.27–1.43 (m, 2H, $\text{CH}_2\text{CH}_2\text{N}$), 3.27–3.38 (m, 1H, CH_A , CH_2N), 3.87–4.01 (m, 1H, CH_B , CH_2N), 5.27 (d, 1H, $^2J_{HP} = 11.0$, C_1H), 6.97 (d, 1H, $J_{HH} = 7.4$, ArH), 7.31–7.43 (m, 2H, ArH), 7.44–7.63 (m, 2H, ArH), 7.77–7.88 (m, 2H, ArH); ^{13}C NMR (CDCl_3) δ 13.7 ($\text{CH}_3(\text{CH}_2)_3\text{N}$), 19.9 ($\text{CH}_2(\text{CH}_2)_2\text{N}$), 26.0 ($\text{C}(\text{CH}_3)_3$), 29.6 ($\text{CH}_2\text{CH}_2\text{N}$), 35.3 (d, $^1J_{P-C} = 62.0$, $\text{C}(\text{CH}_3)_3$), 42.4 (CH_2N), 61.8 (d, $^1J_{CP} = 60.0$, C_1), 124.1 (d, $J_{CP} = 1.1$, C_4), 125.3 (d, $^3J_{CP} = 1.8$, C_7), 128.4 (d, $^3J_{CP} = 10.7$, C_3'), 129.0 (d, $J_{CP} = 1.5$, C_5), 131.1 (d, $J_{CP} = 2.0$, C_6), 131.9 (d, $^2J_{CP} = 8.1$, C_2'), 132.35 (d, $J_{CP} = 2.0$, C_4'), 132.39 (d, $J_{CP} = 2.7$, C_{3a}), 132.9 (d, $^1J_{CP} = 39.1$, C_1'), 140.0 (d, $^2J_{CP} = 2.1$, C_{7a}), 169.5 (d, $^3J_{CP} = 1.3$, C_3); ^{31}P (CDCl_3) δ 45.8; **Isomer B:** ^1H NMR (CDCl_3) δ 0.94 (t, $J_{HH} = 7.2$, 3H, $\text{CH}_3(\text{CH}_2)_3\text{N}$), 1.14 (d, 9H, $^3J_{HP} = 15.0$, $\text{C}(\text{CH}_3)_3$), 1.27–1.43 (m, 2H, $\text{CH}_2(\text{CH}_2)_2\text{N}$), 1.60–1.77 (m, 2H, $\text{CH}_2\text{CH}_2\text{N}$), 3.87–4.01 (m, 1H, CH_A , CH_2N), 4.23–4.36 (m, 1H, CH_B , CH_2N), 5.15 (d, 1H, $^2J_{HP} = 11.9$, C_1H), 7.31–7.43 (m, 2H, ArH), 7.44–7.63 (m, 2H, ArH), 7.77–7.88 (m, 2H, ArH), 8.02 (d, 1H, $J_{HH} = 7.5$, ArH); ^{13}C NMR (CDCl_3) δ 13.8 ($\text{CH}_3(\text{CH}_2)_3\text{N}$), 20.1 ($\text{CH}_2(\text{CH}_2)_2\text{N}$), 25.9 ($\text{C}(\text{CH}_3)_3$), 30.0 ($\text{CH}_2\text{CH}_2\text{N}$), 35.9 (d, $^1J_{P-C} = 60.8$, $\text{C}(\text{CH}_3)_3$), 43.2 (CH_2N), 62.4 (d, $^1J_{CP} = 62.0$, C_1), 123.9 (d, $J_{CP} = 1.3$, C_4), 124.3 (d, $^3J_{CP} = 2.2$, C_7), 128.5 (d, $^3J_{CP} = 10.5$, C_3'), 128.7 (d, $J_{CP} = 1.6$, C_5), 129.8 (d, $^1J_{CP} = 45.1$, C_1'), 131.0 (d, $J_{CP} = 1.9$, C_6), 132.3 (d, $^2J_{CP} = 7.8$, C_2'), 132.40 (d, $J_{CP} = 1.9$, C_4'), 132.41 (d, $J_{CP} = 2.9$, C_{3a}), 139.9 (d, $^2J_{CP} = 1.7$, C_{7a}), 169.3 (d, $^3J_{CP} = 1.9$, C_3); ^{31}P (CDCl_3) δ 44.4; $[\text{M}+\text{H}]^+_{\text{found}} = 370.1946$, $\text{C}_{22}\text{H}_{29}\text{NO}_2\text{P}$ requires 370.1930.

2-Methylphenyl(phenyl) (2-butyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (13)

Yield: 96% (0.39 g), *dr*: 40:60, light yellow crystals; Mp: 118–120 °C; characterized as a mixture, **Isomer A:** ^1H NMR (CDCl_3) δ 0.80 (t, $J_{HH} = 7.3$, 3H, $\text{CH}_3(\text{CH}_2)_3\text{N}$), 0.85–0.96 (m,

1H, CH_A, CH₂(CH₂)₂N), 1.04–1.17 (m, 1H, CH_B, CH₂(CH₂)₂N), 1.37–1.55 (m, 2H, CH₂CH₂N), 2.64–2.77 (m, 1H, CH_A, CH₂N), 2.89 (s, 3H, C₂'CH₃), 3.86–4.02 (m, 1H, CH_B, CH₂N), 5.55 (d, 1H, $^2J_{\text{HP}} = 10.1$, C₁H), 6.63 (d, 1H, $J_{\text{HH}} = 7.7$, ArH), 7.08–7.22 (m, 2H, ArH), 7.23–7.39 (m, 4H, ArH), 7.39–7.60 (m, 5H, ArH), 7.63 (d, 1H, $J_{\text{HH}} = 6.9$, ArH); ^{13}C NMR (CDCl₃) δ 13.6 (CH₃(CH₂)₃N), 19.9 (CH₂(CH₂)₂N), 21.8 (d, $^3J_{\text{CP}} = 8.0$, C₂'CH₃), 30.2 (CH₂CH₂N), 41.6 (CH₂N), 60.5 (d, $^1J_{\text{CP}} = 71.5$, C₁), 123.4 (d, $J_{\text{CP}} = 1.5$, C₄), 123.6 (d, $^3J_{\text{CP}} = 2.1$, C₇), 125.1 (d, $^1J_{\text{CP}} = 94.6$, C₁’)*, 125.4 (d, $^3J_{\text{CP}} = 12.5$, C₅’), 128.3 (d, $^3J_{\text{CP}} = 11.6$, C₃’)*, 128.5 (d, $J_{\text{CP}} = 1.6$, C₅), 130.9 (d, $^1J_{\text{CP}} = 95.6$, C₁’)*, 131.1 (d, $J_{\text{CP}} = 1.7$, C₆), 131.8 (d, $^2J_{\text{CP}} = 10.0$, C₂’)*, 132.3 (d, $^3J_{\text{CP}} = 11.9$, C₃’)*, 132.4 (d, $J_{\text{CP}} = 3.2$, C₄’)*, 132.7 (d, $^2J_{\text{CP}} = 9.5$, C₆’)*, 132.8 (d, $J_{\text{CP}} = 2.3$, C₄’)*, 133.3 (d, $J_{\text{CP}} = 1.3$, C_{3a}), 138.5 (d, $^2J_{\text{CP}} = 1.7$, C_{7a}), 142.3 (d, $^2J_{\text{CP}} = 8.0$, C₂’), 168.3 (d, $^3J_{\text{CP}} = 1.9$, C₃’), *tentative assignments; ^{31}P (CDCl₃) δ 31.1; **Isomer B:** ^1H NMR (CDCl₃) δ 0.90 (t, $J_{\text{HH}} = 7.2$, 3H, CH₃(CH₂)₃N), 1.19–1.37 (m, 2H, CH₂(CH₂)₂N), 1.56–1.78 (m, 2H, CH₂CH₂N), 2.91 (s, 3H, C₂'CH₃), 3.56–3.68 (m, 1H, CH_A, CH₂N), 3.86–4.02 (m, 1H, CH_B, CH₂N), 5.64 (d, 1H, $J_{\text{HP}} = 11.6$, C₁H), 7.08–7.22 (m, 3H, ArH), 7.23–7.39 (m, 4H, ArH), 7.39–7.60 (m, 5H, ArH), 7.75 (d, 1H, $J_{\text{HH}} = 7.5$, ArH); ^{13}C NMR (CDCl₃) δ 13.8 (CH₃(CH₂)₃N), 20.0 (CH₂(CH₂)₂N), 21.8 (d, $^3J_{\text{CP}} = 8.2$, C₂'CH₃), 30.0 (CH₂CH₂N), 42.0 (CH₂N), 61.5 (d, $^1J_{\text{CP}} = 70.8$, C₁), 123.9 (d, $J_{\text{CP}} = 1.0$, C₄), 124.5 (d, $^3J_{\text{CP}} = 2.3$, C₇), 125.8 (d, $^3J_{\text{CP}} = 12.5$, C₅’), 126.8 (d, $^1J_{\text{CP}} = 99.9$, C₁’)*, 128.3 (d, $^3J_{\text{CP}} = 11.9$, C₃’)*, 128.8 (d, $J_{\text{CP}} = 2.2$, C₅), 130.9 (d, $^1J_{\text{CP}} = 96.3$, C₁’)*, 131.2 (d, $J_{\text{CP}} = 2.1$, C₆), 131.9 (d, $^2J_{\text{CP}} = 9.8$, C₂’)*, 132.58 (d, $^3J_{\text{CP}} = 11.3$, C₃’)*, 132.60 (d, $J_{\text{CP}} = 1.8$, C₄’)*, 132.7 (d, $^2J_{\text{CP}} = 9.9$, C₆’)*, 133.3 (d, $J_{\text{CP}} = 3.3$, C₄’)*, 133.6 (d, $J_{\text{CP}} = 1.6$, C_{3a}), 139.7 (d, $^2J_{\text{CP}} = 0.9$, C_{7a}), 143.7 (d, $^2J_{\text{CP}} = 7.8$, C₂’), 169.3 (d, $^3J_{\text{CP}} = 1.3$, C₃’), *tentative assignments; ^{31}P (CDCl₃) δ 34.7; [M+H]⁺_{found} = 404.1790, C₂₅H₂₇NO₂P requires 404.1773.

2-Methoxyphenyl(phenyl) (2-butyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (14)

Yield: 95% (0.40 g), *dr*: 45:55 white crystals; Mp: 133–135 °C; characterized as a mixture, **Isomer A:** ^1H NMR (CDCl₃) δ 0.79 (t, $J_{\text{HH}} = 7.2$, 3H, CH₃(CH₂)₃N), 0.99–1.15 (m, 2H, CH₂(CH₂)₂N), 1.36–1.44 (m, 1H, CH_A, CH₂CH₂N), 1.46–1.53 (m, 1H, CH_B, CH₂CH₂N), 2.66–2.74 (m, 1H, CH_A, CH₂N), 3.86–3.94 (m, 1H, CH_B, CH₂N), 3.96 (s, 3H, CH₃O), 5.86 (d, 1H, $^2J_{\text{HP}} = 10.4$, C₁H), 6.99 (t, 1H, $J_{\text{HH}} = 6.9$, ArH), 7.03–7.12 (m, 1H, ArH), 7.15–7.22 (m, 1H, ArH), 7.22–7.31 (m, 4H, ArH), 7.32–7.42 (m, 2H, ArH), 7.45 (t, 1H, $J_{\text{HH}} = 7.6$, ArH), 7.52–7.59 (m, 2H, ArH), 7.68 (d, 1H, $J_{\text{HH}} = 7.6$, ArH); ^{13}C NMR (CDCl₃) δ 13.7 (CH₃(CH₂)₃N), 20.0 (CH₂(CH₂)₂N), 30.2 (CH₂CH₂N), 41.4 (CH₂N), 55.7 (CH₃O), 60.1 (d,

$^1J_{CP} = 74.0$, C₁), 111.2 (d, $^2J_{CP} = 6.6$, C₆'), 119.5 (d, $^1J_{CP} = 94.6$, C₁'')*, 121.3 (d, $^3J_{CP} = 11.5$, C₅'), 123.2 (d, $J_{CP} = 1.1$, C₄), 123.4 (d, $^3J_{CP} = 2.2$, C₇), 126.6 (d, $^1J_{CP} = 99.8$, C₁')*, 128.0 (d, $^3J_{CP} = 11.7$, C₃''), 128.2 (d, $J_{CP} = 0.8$, C₅), 131.0 (d, $J_{CP} = 2.0$, C₆), 131.7 (d, $^2J_{CP} = 8.3$, C₂''), 132.2 (d, $J_{CP} = 3.0$, C₄'')*, 133.5 (d, $J_{CP} = 3.3$, C₄')*, 134.3 (d, $^3J_{CP} = 6.9$, C₃'), 135.0 (d, $J_{CP} = 2.2$, C_{3a}), 139.7 (d, $^2J_{CP} = 3.3$, C_{7a}), 161.5 (d, $^2J_{CP} = 4.1$, C₂'), 168.2 (d, $^3J_{CP} = 1.3$, C₃), *tentative assignments; ^{31}P (CDCl₃) δ 29.3; **Isomer B:** 1H NMR (CDCl₃) δ 0.87 (t, $J_{HH} = 7.3$, 3H, CH₃(CH₂)₃N), 1.18–1.28 (m, 2H, CH₂(CH₂)₂N), 1.54–1.62 (m, 1H, CH_A, CH₂CH₂N), 1.63–1.72 (m, 1H, CH_B, CH₂CH₂N), 3.48–3.56 (m, 1H, CH_A, CH₂N), 3.86–3.94 (m, 1H, CH_B, CH₂N), 4.05 (s, 3H, CH₃O), 5.92 (d, 1H, $^2J_{HP} = 11.9$, C₁H), 6.84 (d, 1H, $J_{HH} = 7.6$, ArH), 6.92 (t, 1H, $J_{HH} = 7.2$, ArH), 7.03–7.12 (m, 1H, ArH), 7.15–7.22 (m, 1H, ArH), 7.22–7.31 (m, 4H, ArH), 7.32–7.42 (m, 2H, ArH), 7.52–7.59 (m, 3H, ArH); ^{13}C NMR (CDCl₃) δ 13.8 (CH₃(CH₂)₃N), 20.0 (CH₂(CH₂)₂N), 30.0 (CH₂CH₂N), 41.9 (CH₂N), 56.0 (CH₃O), 61.5 (d, $^1J_{CP} = 72.9$, C₁), 111.3 (d, $^2J_{CP} = 6.8$, C₆'), 119.5 (d, $^1J_{CP} = 94.7$, C₁'')*, 121.2 (d, $^3J_{CP} = 11.6$, C₅'), 123.7 (d, $J_{CP} = 1.1$, C₄), 124.4 (d, $^3J_{CP} = 2.1$, C₇), 126.7 (d, $^1J_{CP} = 99.9$, C₁')*, 128.0 (d, $^3J_{CP} = 11.7$, C₃''), 128.5 (d, $J_{CP} = 1.2$, C₅), 130.9 (d, $J_{CP} = 2.2$, C₆), 132.0 (d, $^2J_{CP} = 8.6$, C₂''), 132.5 (d, $J_{CP} = 2.9$, C₄'')*, 133.5 (d, $J_{CP} = 3.0$, C₄')*, 134.3 (d, $^3J_{CP} = 7.4$, C₃'), 134.8 (d, $J_{CP} = 1.9$, C_{3a}), 138.5 (d, $^2J_{CP} = 2.4$, C_{7a}), 161.2 (d, $^2J_{CP} = 3.5$, C₂'), 169.3 (d, $^3J_{CP} = 1.8$, C₃), *tentative assignments; ^{31}P (CDCl₃) δ 33.7; [M+H]⁺_{found} = 420.1739, C₂₅H₂₇NO₃P requires 420.1723.

2-Trifluoromethylphenyl(phenyl) (2-butyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (15)

Yield: 97% (0.44 g), *dr*: 35:65, colourless oil; characterized as a mixture, **Isomer A:** 1H NMR (CDCl₃) δ 0.77 (t, $J_{HH} = 7.4$, 3H, CH₃(CH₂)₃N), 0.97–1.17 (m, 2H, CH₂(CH₂)₂N), 1.19–1.39 (m, 1H, CH_A, CH₂CH₂N), 1.38–1.52 (m, 1H, CH_B, CH₂CH₂N), 2.54–2.64 (m, 1H, CH_A, CH₂N), 3.84–3.95 (m, 1H, CH_B, CH₂N), 5.63 (d, 1H, $^2J_{HP} = 10.7$, C₁H), 6.57 (d, 1H, $J_{HH} = 7.6$, ArH), 7.13 (d, 1H, $J_{HH} = 6.6$, ArH), 7.29–7.41 (m, 3H, ArH), 7.42–7.60 (m, 4H, ArH), 7.67 (d, 1H, $J_{HH} = 7.2$, ArH), 7.74 (d, 1H, $J_{HH} = 6.9$, ArH), 7.98–8.02 (m, 4H, ArH); ^{13}C NMR (CDCl₃) δ 13.5 (CH₃(CH₂)₃N), 19.9 (CH₂(CH₂)₂N), 29.7 (CH₂CH₂N), 41.9 (CH₂N), 61.7 (dd, $^1J_{CP} = 74.9$, $J_{CF} = 2.5$, C₁), 123.6 (d, $^3J_{CP} = 0.9$, C₇), 124.5 (d, $J_{CP} = 2.3$, C₄), 126.6 (d, $^1J_{CP} = 103.6$, C₁''), 127.2 ($^1J_{CP} = 100.7$, C₁'), 128.5 (d, $^3J_{CP} = 11.6$, C₃''), 128.7 (m, C₃'), 129.1 (d, $J_{CP} = 1.8$, C₅), 130.3 (dd, $^3J_{CP} = 2.1$, $^1J_{CF} = 235.0$, CF₃), 131.6 (dd, $^2J_{CP} = 11.3$, $^2J_{CF} = 36.4$, C₂'), 131.5 (d, $J_{CP} = 2.1$, C₆), 131.9 (d, $^2J_{CP} = 8.2$, C₂''), 132.7 (d, $J_{CP} = 4.0$, C₄''),

132.9 (d, $J_{CP} = 3.1$, C_{4'}), 133.0 (d, $^2J_{CP} = 6.6$, C_{6'}), 133.6 (d, $J_{CP} = 3.2$, C_{3a}), 134.2 (d, $^3J_{CP} = 10.5$, C_{5'}), 139.7 (d, $^2J_{CP} = 2.4$, C_{7a}), 169.6 (d, $^3J_{CP} = 2.1$, C₃); ^{31}P (CDCl₃) δ 28.4; **Isomer B:** 1H NMR (CDCl₃) δ 0.89 (t, $J_{HH} = 7.2$, 3H, CH₃(CH₂)₃N), 1.19–1.31 (m, 2H, CH₂(CH₂)₂N), 1.54–1.66 (m, 1H, CH_A, CH₂CH₂N), 1.66–1.76 (m, 1H, CH_B, CH₂CH₂N), 3.47–3.56 (m, 1H, CH_A, CH₂N), 3.84–3.95 (m, 1H, CH_B, CH₂N), 5.72 (d, 1H, $^2J_{HP} = 12.6$, C_{1H}), 7.11 (d, 1H, $J_{HH} = 7.6$, ArH), 7.28 (d, 1H, $J_{HH} = 5.1$, ArH), 7.29–7.41 (m, 3H, ArH), 7.42–7.60 (m, 4H, ArH), 7.80 (d, 1H, $J_{HH} = 7.3$, ArH), 7.74 (d, 1H, $J_{HH} = 6.9$, ArH), 8.02–8.07 (m, 4H, ArH); ^{13}C NMR (CDCl₃) δ 13.7 (CH₃(CH₂)₃N), 19.9 (CH₂(CH₂)₂N), 29.9 (CH₂CH₂N), 42.1 (CH₂N), 61.9 (dd, $^1J_{CP} = 72.6$, $J_{CF} = 3.0$, C₁), 123.5 (d, $J_{CP} = 2.1$, C₄), 124.2 (d, $^3J_{CP} = 0.9$, C₇), 126.6 (d, $^1J_{CP} = 103.8$, C_{1''}), 127.2 ($^1J_{CP} = 100.9$, C₁), 128.5 (d, $^3J_{CP} = 11.6$, C_{3''}), 128.7 (m, C_{3'}), 129.1 (d, $J_{CP} = 1.4$, C₅), 130.3 (dd, $^3J_{CP} = 2.1$, $^1J_{CF} = 239.8$, CF₃), 131.2 (d, $J_{CP} = 1.8$, C₆), 131.6 (dd, $^2J_{CP} = 11.3$, $^2J_{CF} = 37.2$, C_{2'}), 132.2 (d, $^2J_{CP} = 7.8$, C_{2''}), 132.7 (d, $J_{CP} = 3.3$, C_{4''}), 133.00 (d, $^2J_{CP} = 6.6$, C_{6'}), 133.03 (d, $J_{CP} = 3.1$, C_{4'}), 133.6 (d, $J_{CP} = 1.8$, C_{3a}), 134.5 (d, $^3J_{CP} = 10.5$, C_{5'}), 138.2 (d, $^2J_{CP} = 1.8$, C_{7a}), 168.3 (d, $^3J_{CP} = 2.1$, C₃); ^{31}P (CDCl₃) δ 32.7; [M+H]⁺_{found} = 458.1504, C₂₅H₂₄NO₂F₃P requires 458.1491.

3-Trifluoromethylphenyl(phenyl) (2-butyl-3-oxo-2,3-dihydro-2H-isoindol-1-yl)phosphine oxide (16)

Yield: 96% (0.44 g), *dr*: 50:55, colourless oil; characterized as a mixture, **Isomer A:** 1H NMR (CDCl₃) δ 0.77 (t, $J_{HH} = 7.5$, 3H, CH₃(CH₂)₃N), 1.01–1.23 (m, 2H, CH₂(CH₂)₂N), 1.37–1.64 (m, 2H, CH₂CH₂N), 2.94–3.02 (m, 1H, CH_A, CH₂N), 3.83–3.97 (m, 1H, CH_B, CH₂N), 5.43 (d, 1H, $^2J_{HP} = 10.5$, C_{1H}), 6.70 (d, 1H, $J_{HH} = 7.6$, ArH), 7.24–7.65 (m, 10H, ArH), 7.67–7.78 (m, 2H, ArH); ^{13}C NMR (CDCl₃) δ 13.6 (CH₃(CH₂)₃N), 19.8 (CH₂(CH₂)₂N), 29.9 (CH₂CH₂N), 41.7 (CH₂N), 60.7 (d, $^1J_{CP} = 73.8$, C₁), 123.2 (dd, $J_{CP} = 1.6$, $^1J_{CF} = 274.3$, CF₃), 123.76 (d, $^3J_{CP} = 2.5$, C₇), 123.83 (d, $J_{CP} = 0.9$, C₄), 127.7 (d, $^1J_{CP} = 98.6$, C_{1''})*, 128.4 (d, $^1J_{CP} = 97.3$, C_{1''})*, 128.6 (m, C_{2'}), 128.7 (dd, $^3J_{CP} = 9.4$, $^2J_{CF} = 30.6$, C_{3'}), 129.00 (d, $^3J_{CP} = 11.8$, C_{3''}), 129.04 (d, $^3J_{CP} = 13.3$, C_{5'}), 129.3 (d, $J_{CP} = 2.8$, C₅), 129.4 (d, $J_{CP} = 4.0$, C_{4''}), 131.3 (d, $J_{CP} = 1.9$, C₆), 131.4 (d, $^2J_{CP} = 9.1$, C_{2''}), 132.6 (d, $J_{CP} = 3.1$, C_{4'}), 133.3 (d, $J_{CP} = 2.8$, C_{3a}), 134.5 (d, $^2J_{CP} = 8.6$, C_{6'}), 137.9 (d, $^2J_{CP} = 2.0$, C_{7a}), 168.3 (d, $^3J_{CP} = 1.6$, C₃), *tentative assignments; ^{31}P (CDCl₃) δ 29.2; **Isomer B:** 1H NMR (CDCl₃) δ 0.81 (t, $J_{HH} = 7.3$, 3H, CH₃(CH₂)₃N), 1.01–1.23 (m, 2H, CH₂(CH₂)₂N), 1.37–1.64 (m, 2H, CH₂CH₂N), 3.29–3.97 (m, 1H, CH_A, CH₂N), 3.83–3.97 (m, 1H, CH_B, CH₂N), 5.47 (d, 1H, $^2J_{HP} = 10.9$, C_{1H}), 6.98 (d, 1H, $J_{HH} = 7.2$, ArH), 7.24–7.65 (m, 10H, ArH), 7.67–7.78 (m, 1H, ArH), 7.88 (d, 1H,

$J_{HH} = 11.3$, ArH); ^{13}C NMR (CDCl_3) δ 13.6 ($\text{CH}_3(\text{CH}_2)_3\text{N}$), 19.9 ($\text{CH}_2(\text{CH}_2)_2\text{N}$), 30.0 ($\text{CH}_2\text{CH}_2\text{N}$), 42.0 (CH_2N), 61.0 (d, $^1J_{\text{CP}} = 73.8$, C₁), 123.3 (dd, $J_{\text{CP}} = 1.4$, $^1J_{\text{CF}} = 273.1$, CF₃), 123.99 (d, $^3J_{\text{CP}} = 2.5$, C₇), 124.00 (d, $J_{\text{CP}} = 1.4$, C₄), 128.5 (d, $^1J_{\text{CP}} = 98.4$, C₁)*, 128.77 (dd, $^3J_{\text{CP}} = 9.4$, $^2J_{\text{CF}} = 31.3$, C₃'), 128.80 (m, C₂'), 129.00 (d, $^3J_{\text{CP}} = 11.8$, C₃''), 129.2 (d, $^3J_{\text{CP}} = 11.6$, C₅'), 129.51 (d, $J_{\text{CP}} = 3.2$, C₄''), 129.53 (d, $J_{\text{CP}} = 2.9$, C₅), 129.6 (d, $^1J_{\text{CP}} = 96.3$, C₁''), 131.3 (d, $J_{\text{CP}} = 2.2$, C₆), 131.5 (d, $^2J_{\text{CP}} = 9.4$, C₂''), 132.9 (d, $J_{\text{CP}} = 3.2$, C₄'), 133.3 (d, $J_{\text{CP}} = 2.8$, C_{3a}), 135.3 (d, $^2J_{\text{CP}} = 8.4$, C₆'), 138.3 (d, $^2J_{\text{CP}} = 3.2$, C_{7a}), 168.6 (d, $^3J_{\text{CP}} = 1.6$, C₃), *tentative assignments; ^{31}P (CDCl_3) δ 30.5; [M+H]⁺_{found} = 458.1505, $\text{C}_{25}\text{H}_{24}\text{NO}_2\text{F}_3\text{P}$ requires 458.1491.

4-Trifluoromethylphenyl(phenyl) (2-butyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (17)

Yield: 98% (0.45 g), *dr*: 50:50, colourless oil; characterized as a mixture, **Isomer A:** ^1H NMR (CDCl_3) δ 0.82 (t, $J_{HH} = 7.2$, 3H, $\text{CH}_3(\text{CH}_2)_3\text{N}$), 1.06–1.33 (m, 2H, $\text{CH}_2(\text{CH}_2)_2\text{N}$), 1.41–1.69 (m, 2H, $\text{CH}_2\text{CH}_2\text{N}$), 2.95–3.04 (m, 1H, CH_A, CH_2N), 3.89–4.02 (m, 1H, CH_B, CH_2N), 5.48 (d, 1H, $^2J_{\text{HP}} = 11.1$, C₁H), 6.75 (d, 1H, $J_{HH} = 7.1$, ArH), 7.29–7.35 (m, 1H, ArH), 7.40–7.56 (m, 5H, ArH), 7.57–7.73 (m, 5H, ArH) 7.73–7.80 (m, 1H, ArH); ^{13}C NMR (CDCl_3) δ 13.6 ($\text{CH}_3(\text{CH}_2)_3\text{N}$), 19.8 ($\text{CH}_2(\text{CH}_2)_2\text{N}$), 29.9 ($\text{CH}_2\text{CH}_2\text{N}$), 41.7 (CH_2N), 60.7 (d, $^1J_{\text{CP}} = 74.2$, C₁), 123.87 (d, $J_{\text{CP}} = 1.2$, C₄), 123.90 (d, $^3J_{\text{CP}} = 2.4$, C₇), 125.4 (d, $^1J_{\text{CF}} = 261.1$, CF₃), 125.4 (m, C₃'), 129.02 (d, $^3J_{\text{CP}} = 11.5$, C₃''), 129.03 (d, $J_{\text{CP}} = 1.4$, C₅), 131.3 (dd, $J_{\text{CP}} = 1.8$, $^2J_{\text{CF}} = 16.0$, C₄'), 131.47 (d, $J_{\text{CP}} = 2.2$, C₆), 131.49 (d, $^2J_{\text{CP}} = 9.3$, C₂')*, 132.22 (d, $^2J_{\text{CP}} = 9.0$, C₂'')*, 132.26 (d, $J_{\text{CP}} = 1.8$, C_{3a}), 132.27 (d, $^1J_{\text{CP}} = 92.8$, C₁')*, 132.5 (d, $^1J_{\text{CP}} = 104.3$, C₁'')*, 133.4 (d, $J_{\text{CP}} = 1.4$, C₄''), 138.0 (d, $^2J_{\text{CP}} = 2.4$, C_{7a}), 168.4 (d, $^3J_{\text{CP}} = 0.9$, C₃), *tentative assignments; ^{31}P (CDCl_3) δ 29.1; **Isomer B:** ^1H NMR (CDCl_3) δ 0.86 (t, $J_{HH} = 7.1$, 3H, $\text{CH}_3(\text{CH}_2)_3\text{N}$), 1.06–1.33 (m, 2H, $\text{CH}_2(\text{CH}_2)_2\text{N}$), 1.41–1.69 (m, 2H, $\text{CH}_2\text{CH}_2\text{N}$), 3.31–3.40 (m, 1H, CH_A, CH_2N), 3.89–4.02 (m, 1H, CH_B, CH_2N), 5.51 (d, 1H, $^2J_{\text{HP}} = 11.2$, C₁H), 7.14 (d, 1H, $J_{HH} = 6.4$, ArH), 7.35–7.40 (m, 1H, ArH), 7.40–7.56 (m, 5H, ArH), 7.57–7.73 (m, 5H, ArH) 7.73–7.80 (m, 1H, ArH); ^{13}C NMR (CDCl_3) δ 13.7 ($\text{CH}_3(\text{CH}_2)_3\text{N}$), 19.9 ($\text{CH}_2(\text{CH}_2)_2\text{N}$), 30.0 ($\text{CH}_2\text{CH}_2\text{N}$), 42.0 (CH_2N), 61.1 (d, $^1J_{\text{CP}} = 73.7$, C₁), 124.04 (d, $J_{\text{CP}} = 1.4$, C₄), 124.06 (d, $^3J_{\text{CP}} = 1.7$, C₇), 125.4 (d, $^1J_{\text{CF}} = 261.1$, CF₃), 125.4 (m, C₃'), 129.0 (d, $^3J_{\text{CP}} = 11.0$, C₃''), 129.1 (d, $J_{\text{CP}} = 1.4$, C₅), 131.4 (dd, $J_{\text{CP}} = 1.6$, $^2J_{\text{CF}} = 15.4$, C₄'), 131.5 (d, $^2J_{\text{CP}} = 9.3$, C₂')*, 131.6 (d, $J_{\text{CP}} = 1.6$, C₆), 132.3 (d, $^1J_{\text{CP}} = 94.0$, C₁')*, 132.41 (d, $^2J_{\text{CP}} = 9.0$, C₂'')*, 132.44 (d, $J_{\text{CP}} = 2.3$, C_{3a}), 134.5 (d, $^1J_{\text{CP}} = 103.9$, C₁'')*, 133.4 (d, $J_{\text{CP}} = 1.4$, C₄''), 138.4 (d, $^2J_{\text{CP}} = 2.7$, C_{7a}), 168.8 (d, $^3J_{\text{CP}} = 0.9$,

C_3), *tentative assignments; ^{31}P (CDCl_3) δ 30.1; $[\text{M}+\text{H}]^+_{\text{found}} = 458.1506$, $\text{C}_{25}\text{H}_{24}\text{NO}_2\text{F}_3\text{P}$ requires 458.1491.

2-Phenyl-phenyl(phenyl) (2-butyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (18)

Yield: 96% (0.45 g), dr : 40:60, yellow oil; characterized as a mixture, **Isomer A:** ^1H NMR (CDCl_3) δ 0.83 (t, $J_{\text{HH}} = 7.5$, 3H, $\text{CH}_3(\text{CH}_2)_3\text{N}$), 0.98–1.09 (m, 2H, $\text{CH}_2(\text{CH}_2)_2\text{N}$), 1.19–1.36 (m, 2H, $\text{CH}_2\text{CH}_2\text{N}$), 2.47–2.55 (m, 1H, CH_A , CH_2N), 3.65–3.76 (m, 1H, CH_B , CH_2N), 4.47 (d, 1H, $^2J_{\text{HP}} = 12.3$, C_1H), 6.56 (d, 1H, $J_{\text{HH}} = 7.7$, ArH), 7.07–7.19 (m, 2H, ArH), 7.21–7.28 (m, 2H, ArH), 7.28–7.43 (m, 5H, ArH), 7.43–7.49 (m, 2H, ArH), 7.50–7.55 (m, 1H, ArH), 7.55–7.61 (m, 1H, ArH), 7.61–7.66 (m, 1H, ArH), 7.66–7.74 (m, 1H, ArH), 7.78 (d, 1H, $J_{\text{HH}} = 7.8$, ArH), 7.86–8.02 (m, 1H, ArH); ^{13}C NMR (CDCl_3) δ 13.9 ($\text{CH}_3(\text{CH}_2)_3\text{N}$), 20.0 ($\text{CH}_2(\text{CH}_2)_2\text{N}$), 29.6 ($\text{CH}_2\text{CH}_2\text{N}$), 41.9 (CH_2N), 58.6 (d, $^1J_{\text{CP}} = 70.0$, C_1), 123.0 (d, $J_{\text{CP}} = 1.3$, C_4), 123.1 (d, $^3J_{\text{CP}} = 2.4$, C_7), 125.3 (d, $J_{\text{CP}} = 1.8$, ArC), 127.3 (d, $J_{\text{CP}} = 12.1$, ArC), 127.7 (d, $J_{\text{CP}} = 11.6$, ArC), 128.0 (d, $J_{\text{CP}} = 100.2$, ArC), 128.35 (d, $J_{\text{CP}} = 2.0$, C_5), 128.38 (ArC), 128.8 (ArC), 129.7 (d, $J_{\text{CP}} = 1.8$, C_6), 129.8 (ArC), 131.0 (d, $J_{\text{CP}} = 10.1$, ArC), 131.7 (d, $J_{\text{CP}} = 102.7$, ArC), 132.2 (d, $J_{\text{CP}} = 2.6$, C_{3a}), 132.6 (d, $J_{\text{CP}} = 8.7$, ArC), 132.7 (d, $J_{\text{CP}} = 4.7$, ArC), 132.9 (d, $J_{\text{CP}} = 5.0$, ArC), 133.5 (d, $J_{\text{CP}} = 11.9$, ArC), 138.4 (d, $^2J_{\text{CP}} = 1.7$, C_{7a}), 140.2 (d, $J_{\text{CP}} = 3.1$, ArC), 147.4 (d, $J_{\text{CP}} = 8.8$, ArC), 167.9 (d, $^3J_{\text{CP}} = 1.8$, C_3); **Isomer B:** ^1H NMR (CDCl_3) δ 0.86 (t, $J_{\text{HH}} = 7.5$, 3H, $\text{CH}_3(\text{CH}_2)_3\text{N}$), 1.09–1.18 (m, 2H, $\text{CH}_2(\text{CH}_2)_2\text{N}$), 1.19–1.36 (m, 2H, $\text{CH}_2\text{CH}_2\text{N}$), 3.65–3.76 (m, 1H, CH_A , CH_2N), 3.82–3.91 (m, 1H, CH_B , CH_2N), 4.61 (d, 1H, $^2J_{\text{HP}} = 8.7$, C_1H), 6.85 (d, 1H, $J_{\text{HH}} = 11.4$, ArH), 7.07–7.19 (m, 3H, ArH), 7.21–7.28 (m, 2H, ArH), 7.28–7.43 (m, 5H, ArH), 7.43–7.49 (m, 2H, ArH), 7.50–7.55 (m, 1H, ArH), 7.55–7.61 (m, 1H, ArH), 7.61–7.66 (m, 1H, ArH), 7.66–7.74 (m, 1H, ArH), 7.86–8.02 (m, 1H, ArH); ^{13}C NMR (CDCl_3) δ 14.0 ($\text{CH}_3(\text{CH}_2)_3\text{N}$), 20.3 ($\text{CH}_2(\text{CH}_2)_2\text{N}$), 30.5 ($\text{CH}_2\text{CH}_2\text{N}$), 42.1 (CH_2N), 59.2 (d, $^1J_{\text{CP}} = 71.0$, C_1), 124.2 (d, $J_{\text{CP}} = 0.9$, C_4), 124.5 (d, $^3J_{\text{CP}} = 1.0$, C_7), 125.6 (d, $J_{\text{CP}} = 2.3$, ArC), 127.4 (d, $J_{\text{CP}} = 11.9$, ArC), 127.9 (d, $J_{\text{CP}} = 11.4$, ArC), 128.1 (d, $J_{\text{CP}} = 95.5$, ArC), 128.6 (ArC), 128.7 (d, $J_{\text{CP}} = 2.3$, C_5), 128.9 (ArC), 129.8 (ArC), 130.6 (d, $J_{\text{CP}} = 1.4$, C_6), 131.0 (d, $J_{\text{CP}} = 10.0$, ArC), 131.7 (d, $J_{\text{CP}} = 95.0$, ArC), 132.86 (d, $J_{\text{CP}} = 9.0$, ArC), 132.87 (d, $J_{\text{CP}} = 5.0$, ArC), 133.0 (d, $J_{\text{CP}} = 3.8$, ArC), 133.5 (d, $J_{\text{CP}} = 11.6$, ArC), 133.9 (d, $J_{\text{CP}} = 3.2$, C_{3a}), 139.5 (d, $^2J_{\text{CP}} = 4.1$, C_{7a}), 140.7 (d, $J_{\text{CP}} = 3.4$, ArC), 147.5 (d, $J_{\text{CP}} = 9.0$, ArC), 169.3 (d, $^3J_{\text{CP}} = 1.3$, C_3); ^{31}P (CDCl_3) δ 36.2; $[\text{M}+\text{H}]^+_{\text{found}} = 466.1942$, $\text{C}_{30}\text{H}_{29}\text{NO}_2\text{P}$ requires 466.1930.

1-Naphthyl(phenyl) (2-butyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (19)

Yield: 97% (0.43 g), *dr*: 45:55; characterized after separation by column chromatography:

Isomer A: white crystals; Mp: 178–180 °C; ¹H NMR (CDCl₃) δ 0.43 (t, J_{HH} = 6.5, 3H, CH₃(CH₂)₃N), 0.40–0.51 (m, 1H, CH_A, CH₂(CH₂)₂N), 0.64–0.76 (m, 1H, CH_B, CH₂(CH₂)₂N), 0.78–0.90 (m, 1H, CH_A, CH₂CH₂N), 0.90–1.00 (m, 1H, CH_B, CH₂CH₂N), 1.90–1.98 (m, 1H, CH_A, CH₂N), 3.67–3.76 (m, 1H, CH_B, CH₂N), 5.81 (d, 1H, ²J_{HP} = 10.5, C₁H), 7.28–7.37 (m, 3H, ArH), 7.38–7.52 (m, 4H, ArH), 7.57–7.70 (m, 4H, ArH), 7.76 (d, 1H, J_{HH} = 7.8, ArH), 7.92 (d, 1H, J_{HH} = 7.8, ArH), 8.00 (d, 1H, J_{HH} = 8.3, ArH), 8.10 (d, 1H, J_{HH} = 8.2, ArH), 9.26 (d, 1H, J_{HH} = 8.7, ArH); ¹³C NMR (CDCl₃) δ 13.2 (CH₃(CH₂)₃N), 19.5 (CH₂(CH₂)₂N), 29.8 (CH₂CH₂N), 41.1 (CH₂N), 60.3 (d, ¹J_{CP} = 72.1, C₁), 123.4 (d, J_{CP} = 1.7, C₄), 124.59 (d, ³J_{CP} = 14.4, C₃'), 123.64 (d, ³J_{CP} = 2.1, C₇), 125.7 (d, ¹J_{CP} = 100.1, C₁'), 126.4 (d, ²J_{CP} = 4.4, C₂'), 127.1 (C₆'), 127.2 (d, ¹J_{CP} = 93.7, C₁''), 128.1 (C₅'), 128.3 (d, ³J_{CP} = 11.4, C₃''), 128.5 (d, J_{CP} = 1.7, C₅), 129.5 (d, ⁴J_{CP} = 1.7, C₄'), 131.4 (d, J_{CP} = 1.7, C₆), 131.9 (d, J_{CP} = 3.1, C₄''), 131.0 (d, ²J_{CP} = 7.8, C₂''), 132.5 (d, J_{CP} = 2.5, C_{3a}), 132.5 (d, ³J_{CP} = 11.5, C₈'), 134.0 (d, ²J_{CP} = 5.2, C₉'), 134.1 (d, ³J_{CP} = 5.3, C₁₀'), 134.1 (d, J_{CP} = 2.9, C₇'), 140.0 (d, ²J_{CP} = 3.6, C_{7a}), 169.4 (d, ³J_{CP} = 2.3, C₃); ³¹P (CDCl₃) δ 30.4; [M+H]⁺_{found} = 440.1794, C₂₈H₂₇NO₂P requires 440.1773.

Isomer B: colourless oil; ¹H NMR (CDCl₃) δ 0.95 (t, J_{HH} = 7.3, 3H, CH₃(CH₂)₃N), 1.27–1.37 (m, 2H, CH₂(CH₂)₂N), 1.64–1.87 (m, 2H, CH₂CH₂N), 3.72–3.80 (m, 1H, CH_A, CH₂N), 3.94–4.04 (m, 1H, CH_B, CH₂N), 5.83 (d, 1H, ²J_{HP} = 11.8, C₁H), 5.94 (d, 1H, J_{HH} = 7.7, ArH), 7.09–7.21 (m, 3H, ArH), 7.29–7.42 (m, 5H, ArH), 7.52 (t, 1H, J_{HH} = 7.4, ArH), 7.67 (t, 1H, J_{HH} = 7.5, ArH), 7.71–7.80 (m, 2H, ArH), 8.02 (d, 1H, J_{HH} = 8.2, ArH), 8.12 (d, 1H, J_{HH} = 6.4, ArH), 9.22 (d, 1H, J_{HH} = 8.5, ArH); ¹³C NMR (CDCl₃) δ 13.9 (CH₃(CH₂)₃N), 20.1 (CH₂(CH₂)₂N), 30.2 (CH₂CH₂N), 42.1 (CH₂N), 61.9 (d, ¹J_{CP} = 71.2, C₁), 123.5 (d, J_{CP} = 2.0, C₄), 123.8 (d, ³J_{CP} = 0.9, C₇), 124.2 (d, ³J_{CP} = 13.9, C₃'), 126.3 (d, ¹J_{CP} = 99.5, C₁'), 126.5 (d, ¹J_{CP} = 94.7, C₁''), 126.6 (d, ²J_{CP} = 4.1, C₂'), 127.1 (C₆'), 128.3 (C₅'), 128.3 (d, ³J_{CP} = 11.4, C₃''), 128.8 (d, J_{CP} = 1.6, C₅), 129.4 (d, J_{CP} = 0.8, C₄'), 131.0 (d, J_{CP} = 1.5, C₆), 132.3 (d, ²J_{CP} = 8.3, C₂''), 132.5 (d, ³J_{CP} = 11.0, C₈'), 132.9 (d, J_{CP} = 2.9, C_{3a}), 133.3 (d, J_{CP} = 3.3, C₄''), 134.00 (d, J_{CP} = 3.4, C₇'), 134.02 (d, ²J_{CP} = 4.9, C₉'), 134.1 (d, ³J_{CP} = 4.3, C₁₀'), 138.5 (d, ²J_{CP} = 2.1, C_{7a}), 168.3 (d, ³J_{CP} = 1.8, C₃); ³¹P (CDCl₃) δ 34.8; [M+H]⁺_{found} = 440.1788, C₂₈H₂₇NO₂P requires 440.1773.

General procedure for the synthesis of diphenyl (2-butyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine (20)

A mixture of 1.0 mmol (0.39 g) of diphenyl (2-butyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine oxide (**7a**) and 3.0 mmol (0.37 ml) of phenylsilane was irradiated in a sealed tube at 100–140 °C for 2–6 h under N₂ atmosphere in a CEM® Discover Microwave reactor equipped with a pressure controller. The product (**20**) was utilized without work-up in the next reactions.

General procedure for the synthesis of diphenyl (2-butyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine sulphide (21)

A mixture of 1.0 mmol (0.37 g) of diphenyl (2-butyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine (**20**) and 1.2 mmol (0.30 g) of sulphur in 10 mL of degassed dichloromethane was stirred at 25 °C for 12 h under N₂ atmosphere. The product **21** was obtained by column chromatography using silica gel as the absorbent and dichloromethane:methanol (97:3) as the eluent.

Diphenyl (2-butyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine sulphide (21)

Yield: 81% (0.42 g), white crystal; Mp: 184–186 °C; ¹H NMR (CDCl₃) δ 0.79 (t, J_{HH} = 7.3, 3H, CH₃(CH₂)₃N), 0.93–1.16 (m, 2H, CH₂(CH₂)₂N), 1.33–1.56 (m, 2H, CH₂CH₂N), 2.66–2.82 (m, 1H, CH_A, CH₂N), 3.82–3.99 (m, 1H, CH_B, CH₂N), 5.64 (d, 1H, ²J_{HP} = 3.1, C₁H), 7.04 (d, 1H, J_{HH} = 7.7, ArH), 7.20–7.30 (m, 1H, ArH), 7.31–7.62 (m, 7H, ArH), 7.64–7.81 (m, 3H, ArH), 7.82–7.95 (m, 2H, ArH); ¹³C NMR (CDCl₃) δ 13.6 (CH₃(CH₂)₃N), 19.8 (CH₂(CH₂)₂N), 29.8 (CH₂CH₂N), 41.4 (CH₂N), 63.5 (d, ¹J_{CP} = 53.3, C₁), 123.5 (d, J_{CP} = 1.5, C₄), 124.1 (d, ³J_{CP} = 2.7, C₇), 128.0 (d, ¹J_{CP} = 77.5, C₁^I), 128.6 (d, ³J_{CP} = 11.9, C₃^I), 128.86 (d, ³J_{CP} = 12.1, C₃^{II}), 128.87 (d, J_{CP} = 1.7, C₅), 129.8 (d, ¹J_{CP} = 77.9, C₁^{II}), 130.8 (d, J_{CP} = 2.5, C₆), 130.0 (d, ²J_{CP} = 9.9, C₂^I), 132.1 (d, ²J_{CP} = 9.3, C₂^{II}), 132.36 (d, J_{CP} = 3.0, C₄^I), 132.44 (d, J_{CP} = 3.0, C₄^{II}), 132.7 (d, ³J_{CP} = 3.0, C_{3a}), 138.4 (d, ²J_{CP} = 2.8, C_{7a}), 169.0 (d, ³J_{CP} = 0.9, C₃); ³¹P (CDCl₃) δ 44.6; [M+H]⁺_{found} = 406.1338, C₂₄H₂₅NOPS requires 406.1340.

General procedure for the synthesis of [diphenyl (2-butyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine] dichloroplatinum(II) (22)

A mixture of 1.0 mmol (0.37 g) of diphenyl (2-butyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine (**20**) and 0.5 mmol (0.24 g) of bis(benzonitrile)dichloroplatinum in 10 mL of degassed dichloromethane was stirred at 25 °C for 12 h under N₂ atmosphere. The product **22**

was obtained by column chromatography using silica gel as the absorbent and dichloromethane:methanol (97:3) as the eluent.

[Diphenyl (2-butyl-3-oxo-2,3-dihydro-2*H*-isoindol-1-yl)phosphine] dichloroplatinum(II) (22)

Yield: 80% (0.81 g), yellow crystals; characterized as a mixture, **Isomer A:** ^1H NMR (CDCl_3) δ 0.75 (t, $J_{\text{HH}} = 7.6$, 3H, $\text{CH}_3(\text{CH}_2)_3\text{N}$), 0.94–1.11 (m, 2H, $\text{CH}_2(\text{CH}_2)_2\text{N}$), 1.34–1.54 (m, 2H, $\text{CH}_2\text{CH}_2\text{N}$), 3.20–3.31 (m, 1H, CH_{A} , CH_2N), 3.75–4.87 (m, 1H, CH_{B} , CH_2N), 6.67 (d, 1H, $^2J_{\text{HP}} = 9.5$, C_1H), 7.07–7.48 (m, 11H, ArH), 7.52–7.61 (m, 2H, ArH), 7.63–7.75 (m, 1H, ArH); ^{13}C NMR (CDCl_3) δ 13.8 ($\text{CH}_3(\text{CH}_2)_3\text{N}$), 19.9 ($\text{CH}_2(\text{CH}_2)_2\text{N}$), 30.6 ($\text{CH}_2\text{CH}_2\text{N}$), 41.4 (CH_2N), 53.6 (dd, $^1J_{\text{CP}} = 14.7$, $^3J_{\text{CP}} = 1.3$, C_1), 123.4 (d, $J_{\text{CP}} = 1.8$, C_4), 124.7 (d, $^3J_{\text{CP}} = 0.9$, C_7), 128.1 (d, $^3J_{\text{CP}} = 5.5$, $\text{C}_3^{\text{'I}}$), 128.2 (d, $^3J_{\text{CP}} = 5.0$, $\text{C}_3^{\text{'II}}$), 128.5 (d, $J_{\text{CP}} = 2.3$, C_5), 131.2 (dd, $^1J_{\text{CP}} = 69.9$, $^3J_{\text{CP}} = 1.1$, $\text{C}_1^{\text{'I}}$), 131.3 (dd, $^1J_{\text{CP}} = 74.0$, $^3J_{\text{CP}} = 1.1$, $\text{C}_1^{\text{'II}}$), 131.5 (d, $J_{\text{CP}} = 1.9$, C_6), 132.7 (d, $J_{\text{CP}} = 1.4$, $\text{C}_{3\text{a}}$), 133.27 (d, $J_{\text{CP}} = 5.0$, $\text{C}_4^{\text{'I}}$), 133.32 (d, $^3J_{\text{CP}} = 5.1$, $\text{C}_4^{\text{'II}}$), 134.4 (d, $^2J_{\text{CP}} = 5.5$, $\text{C}_2^{\text{'I}}$), 134.5 (d, $^2J_{\text{CP}} = 6.0$, $\text{C}_2^{\text{'II}}$), 140.1 (d, $^2J_{\text{CP}} = 2.8$, $\text{C}_{7\text{a}}$), 168.4 (d, $^3J_{\text{CP}} = 0.9$, C_3); ^{31}P (CDCl_3) δ 21.2 (t, $^1J_{\text{CPt}} = 2519$); **Isomer B:** ^1H NMR (CDCl_3) δ 0.77 (t, $J_{\text{HH}} = 7.7$, 3H, $\text{CH}_3(\text{CH}_2)_3\text{N}$), 0.94–1.11 (m, 2H, $\text{CH}_2(\text{CH}_2)_2\text{N}$), 1.34–1.54 (m, 2H, $\text{CH}_2\text{CH}_2\text{N}$), 3.20–3.31 (m, 1H, CH_{A} , CH_2N), 3.75–4.87 (m, 1H, CH_{B} , CH_2N), 6.67 (d, 1H, $^2J_{\text{HP}} = 9.5$, C_1H), 7.07–7.48 (m, 11H, ArH), 7.52–7.61 (m, 2H, ArH), 7.63–7.75 (m, 1H, ArH); ^{13}C NMR (CDCl_3) δ 13.8 ($\text{CH}_3(\text{CH}_2)_3\text{N}$), 19.9 ($\text{CH}_2(\text{CH}_2)_2\text{N}$), 30.6 ($\text{CH}_2\text{CH}_2\text{N}$), 41.5 (CH_2N), 53.6 (dd, $^1J_{\text{CP}} = 15.1$, $^3J_{\text{CP}} = 1.3$, C_1), 123.4 (d, $J_{\text{CP}} = 1.8$, C_4), 124.7 (d, $^3J_{\text{CP}} = 0.9$, C_7), 128.1 (d, $^3J_{\text{CP}} = 5.5$, $\text{C}_3^{\text{'I}}$), 128.48 (d, $^3J_{\text{CP}} = 5.0$, $\text{C}_3^{\text{'II}}$), 128.51 (d, $J_{\text{CP}} = 2.3$, C_5), 131.2 (dd, $^1J_{\text{CP}} = 69.9$, $^3J_{\text{CP}} = 0.8$, $\text{C}_1^{\text{'I}}$), 131.3 (dd, $^1J_{\text{CP}} = 74.3$, $^3J_{\text{CP}} = 0.9$, $\text{C}_1^{\text{'II}}$), 131.6 (d, $J_{\text{CP}} = 1.4$, C_6), 132.8 (d, $J_{\text{CP}} = 1.4$, $\text{C}_{3\text{a}}$), 133.31 (d, $J_{\text{CP}} = 4.9$, $\text{C}_4^{\text{'I}}$), 133.33 (d, $^3J_{\text{CP}} = 5.1$, $\text{C}_4^{\text{'II}}$), 134.46 (d, $^2J_{\text{CP}} = 5.5$, $\text{C}_2^{\text{'I}}$), 134.50 (d, $^2J_{\text{CP}} = 6.0$, $\text{C}_2^{\text{'II}}$), 140.1 (d, $^2J_{\text{CP}} = 2.8$, $\text{C}_{7\text{a}}$), 168.5 (d, $^3J_{\text{CP}} = 0.9$, C_3); ^{31}P (CDCl_3) δ 21.2 (t, $^1J_{\text{CPt}} = 2519$); $[\text{M}+\text{Na}]^+_{\text{found}} = 1033.2107$, $\text{C}_{48}\text{H}_{48}\text{N}_2\text{O}_2\text{P}_2\text{PtNa}$ requires 1033.2087.

Single crystal X-ray diffraction measurements

Single-crystal X-ray diffraction data of **7a** and **22** were collected on an Agilent Technologies SuperNova Dual diffractometer using Mo-K α radiation ($\lambda = 0.71073 \text{ \AA}$) at 150 K. The data were processed using CrysAlis Pro [S1]. Structure was solved by ShelXT [S2] using intrinsic phasing and refined by a full-matrix least-squares procedure based on F^2 with ShelXL [S3]

using Olex2 program suite [S4]. All the non-hydrogen atoms were refined anisotropically. Hydrogen atoms were readily located in difference Fourier maps, and were subsequently treated as riding atoms in geometrically idealized positions with C–H = 0.95 Å (aromatic), 1.00 Å (methine), 0.99 Å (methylene) or 0.98 Å (methyl), and with $U_{\text{iso}}(\text{H}) = kU_{\text{eq}}(\text{C})$, where $k = 1.5$ for methyl group and 1.2 for all other H atoms. The crystallographic data are listed in Table S1. Deposition Numbers CCDC 2100126 and 2100127 contain the supplementary crystallographic data for this paper. These data are provided free of charge by the joint Cambridge Crystallographic Data Centre and Fachinformationszentrum Karlsruhe Access Structures service www.ccdc.cam.ac.uk/structures.

Table S1: Crystal data and structure refinement for compounds **7a** and **22**.

	7a	22
CCDC number	2100126	2100127
Empirical formula	C ₂₄ H ₂₄ NO ₂ P	C ₄₈ H ₄₈ Cl ₂ N ₂ O ₂ P ₂ Pt
Formula weight	389.41	1012.81
T/K	150.00(10)	150.00(10)
Crystal system	monoclinic	monoclinic
Space group	<i>P</i> 2 ₁ /c	<i>P</i> 2 ₁ /n
a/Å	11.7785(7)	11.3292(4)
b/Å	17.1967(9)	10.3205(3)
c/Å	11.2241(6)	18.9713(8)
α/°	90	90
β/°	115.786(8)	106.630(4)
γ/°	90	90
V/Å³	2047.1(2)	2125.40(14)
Z	4	2
D_{calc}/g cm⁻³	1.264	1.583
μ/mm⁻¹	0.154	3.544
F(000)	824.0	1016.0
Reflections collected	11169	17175
Data/restraints/parameters	4703/0/254	4870/0/260
R_{int}	0.0317	0.0411
GOF, S	1.059	1.062
R₁, wR₂ [$I \geq 2\sigma(I)$]	0.0429, 0.1008	0.0260, 0.0559
R₁, wR₂ [all data]	0.0597, 0.1157	0.0361, 0.0614
Δρ_{min}, Δρ_{max} [e Å⁻³]	0.31/−0.31	0.93/−1.23

Table S2. Hydrogen bond geometry in **7a** and **22**.

D–H···A	D–H (Å)	H···A (Å)	D···A (Å)	D–H···A (°)	Symmetry code
7a					
C1–H1···O1	1.00	2.18	3.1622(18)	166.7	$x, \frac{1}{2} - y, -\frac{1}{2} + z$
C15–H15···O2	0.95	2.54	3.256(2)	132.1	$1 - x, -\frac{1}{2} + y, 1\frac{1}{2} - z$
C22–H22···O2	0.95	2.32	3.248(2)	164.1	$2 - x, -\frac{1}{2} + y, 1\frac{1}{2} - z$
C11–H11b···Cg4	0.99	2.99	3.843(2)	145	x, y, z
C16–H16···Cg4	0.95	2.74	3.616(2)	154	$1 - x, -y, 1 - z$
22					
C21–H21···O1	0.95	2.48	3.139(4)	127	$\frac{1}{2} + x, 1\frac{1}{2} - y, \frac{1}{2} + z$
C24–H24···Cg2	0.95	2.74	3.616(2)	154	$-x, 1 - y, 1 - z$

Cg4 (for **7a**) and *Cg2* (for **22**) are C19–C24 and C2–C7 ring centroids, respectively.

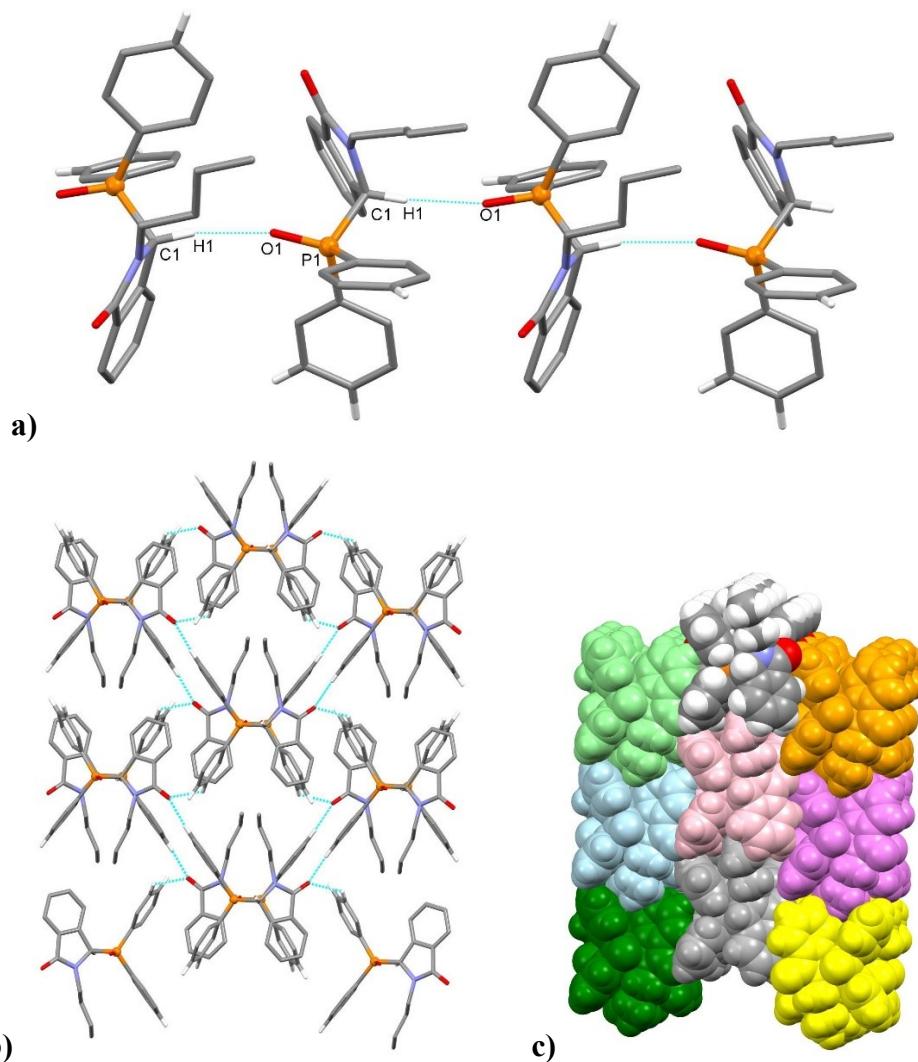


Figure S1. (a) Hydrogen-bonded chain formation in **7a** via C1–H1···O1=Pi interactions along *c*-axis. (b) Hydrogen-bonded network along *ab*-plane formed by C15–H15···O2 and C22–H22···O2.

C22–H22···O2 interactions. (c) Packing of chains (arbitrary colors). Blue dashed lines indicate hydrogen bonds.

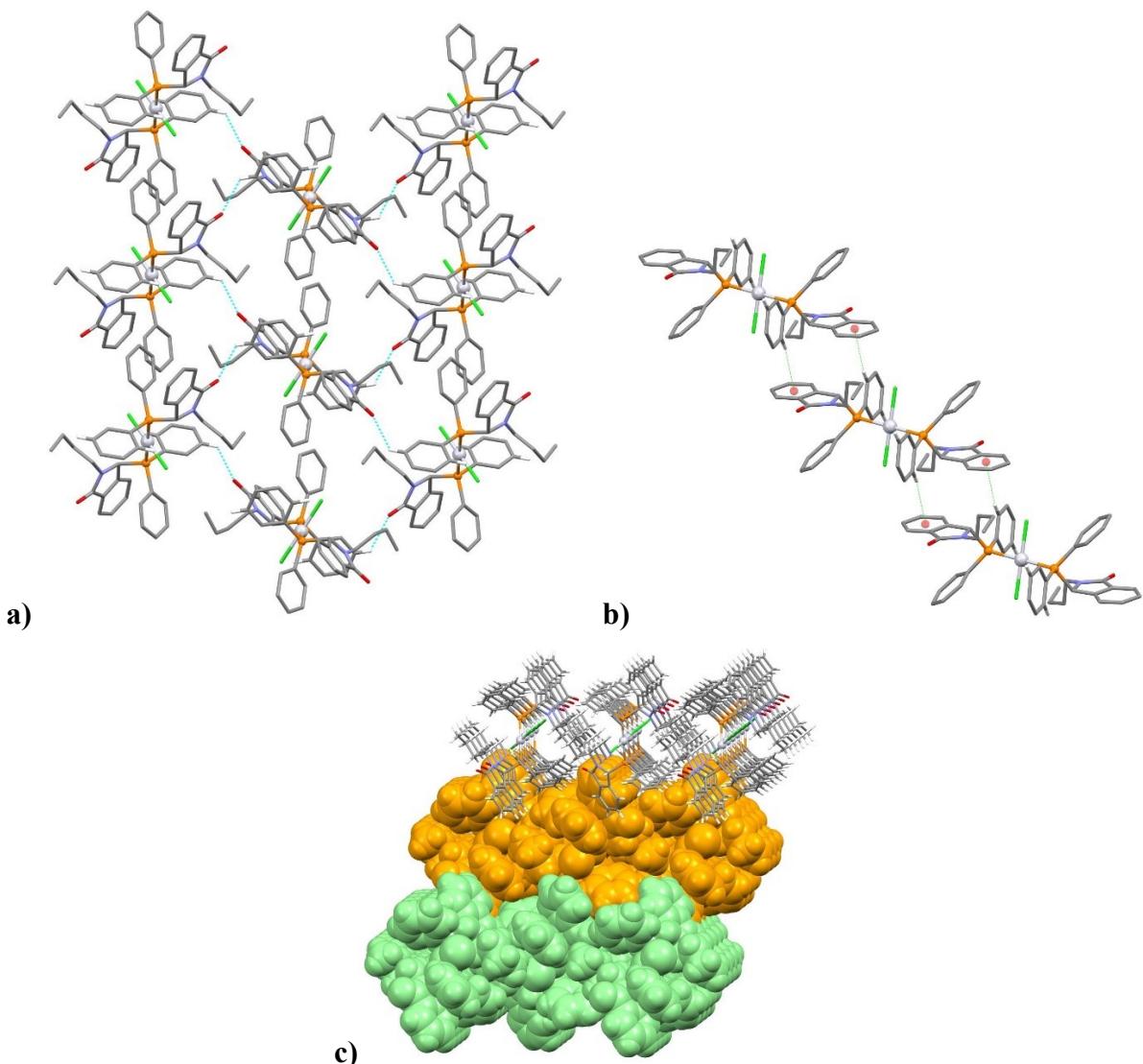
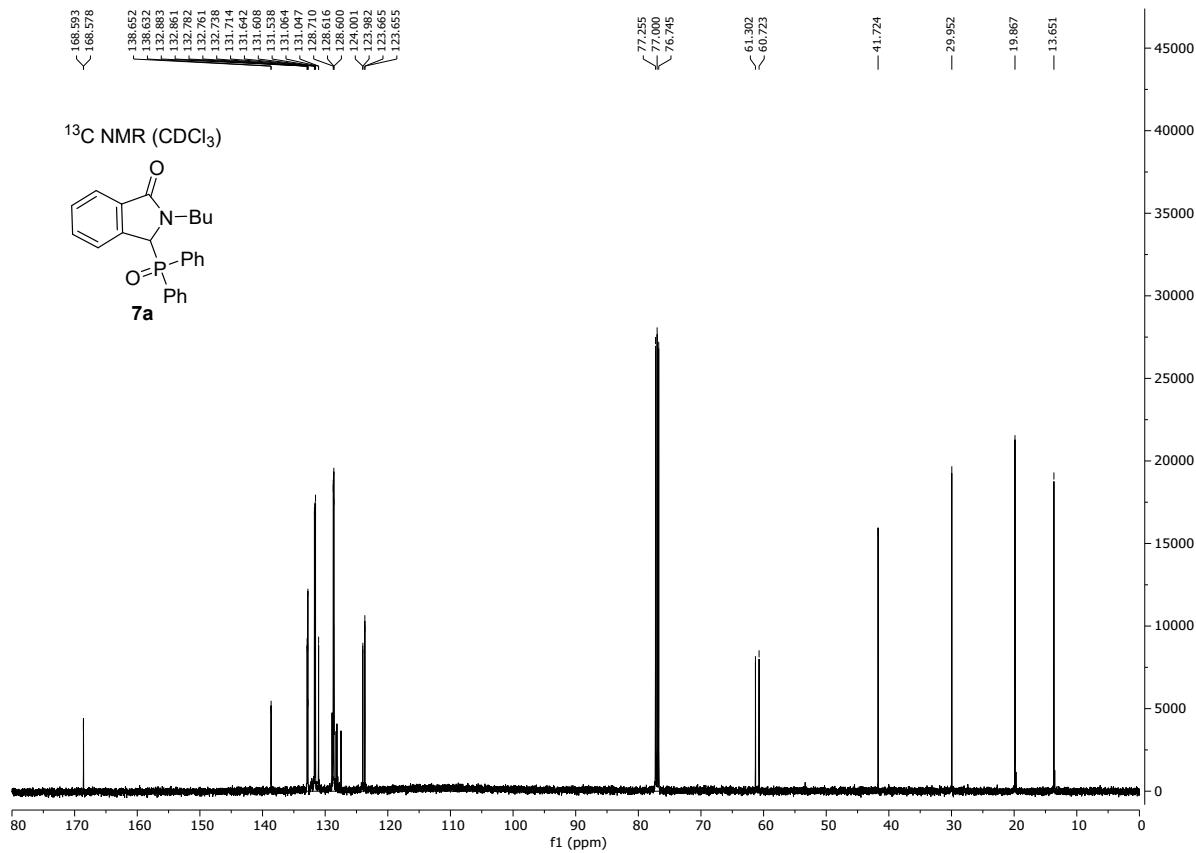
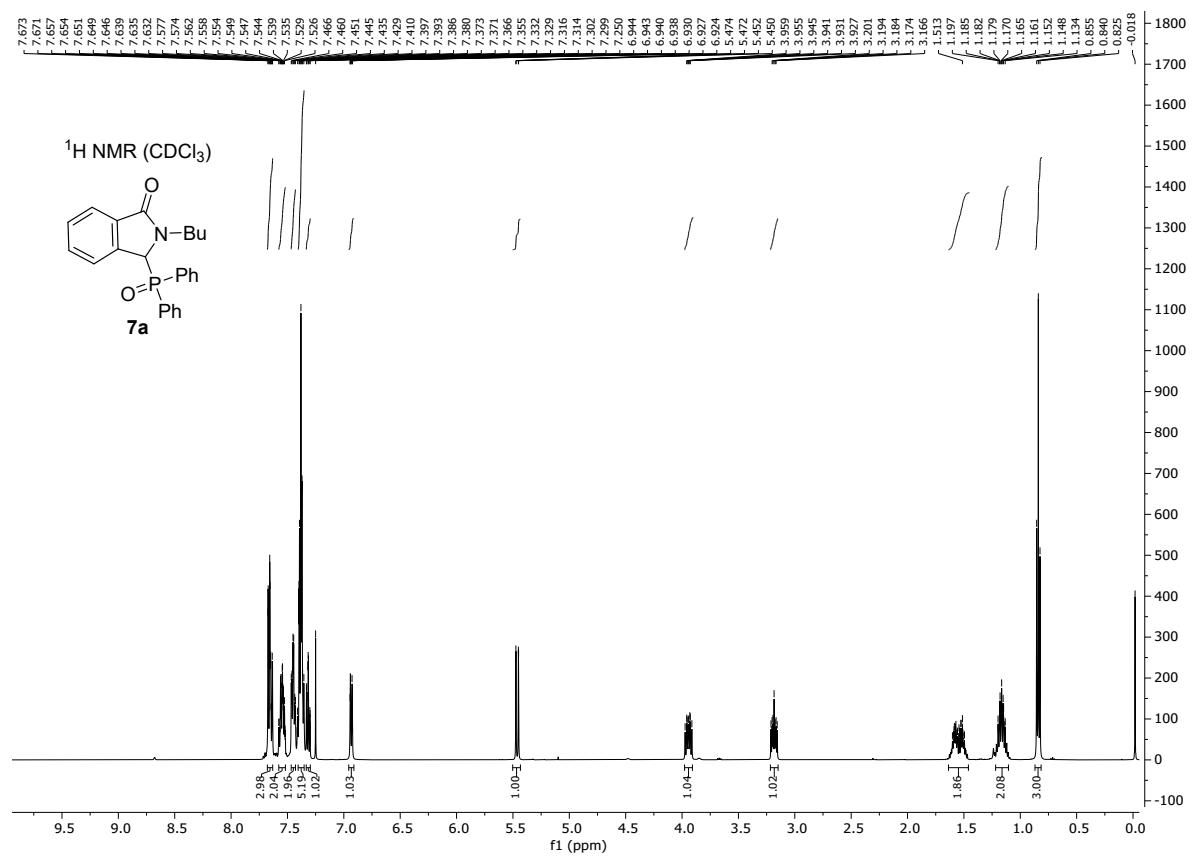
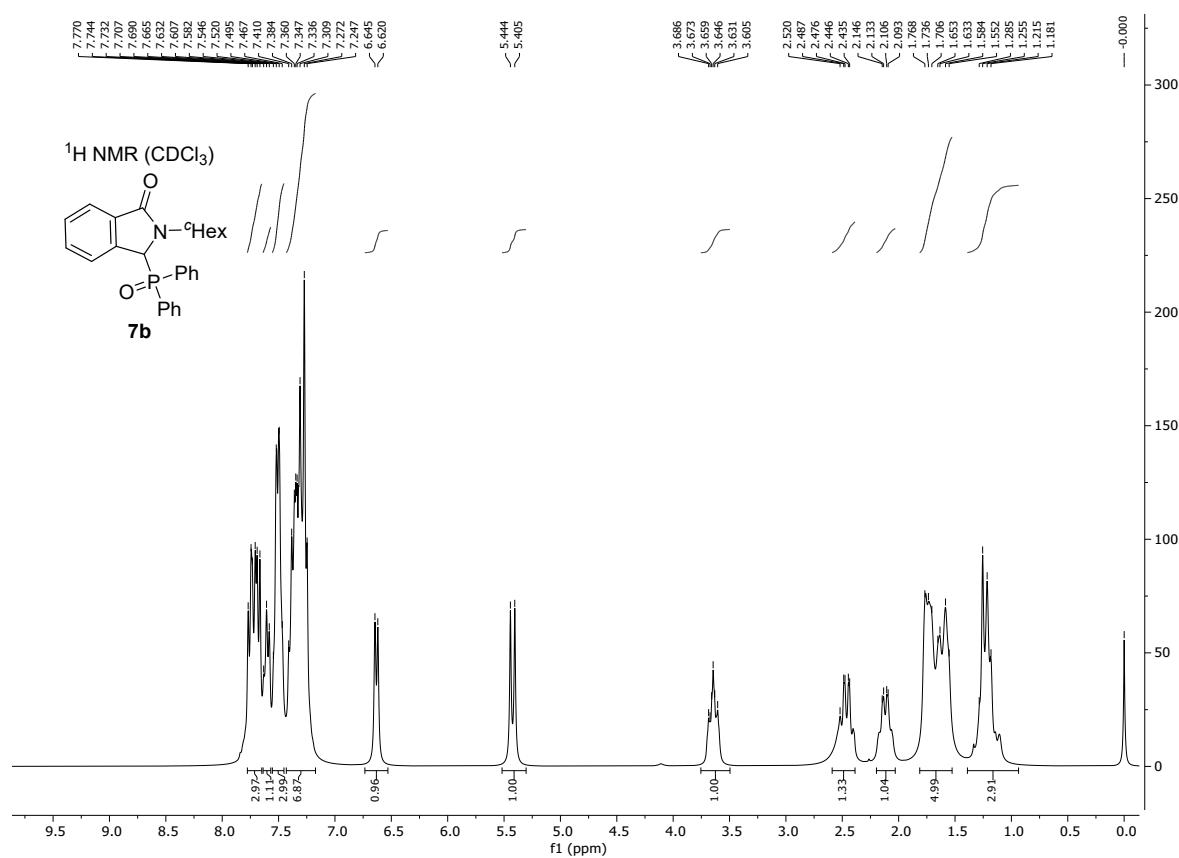
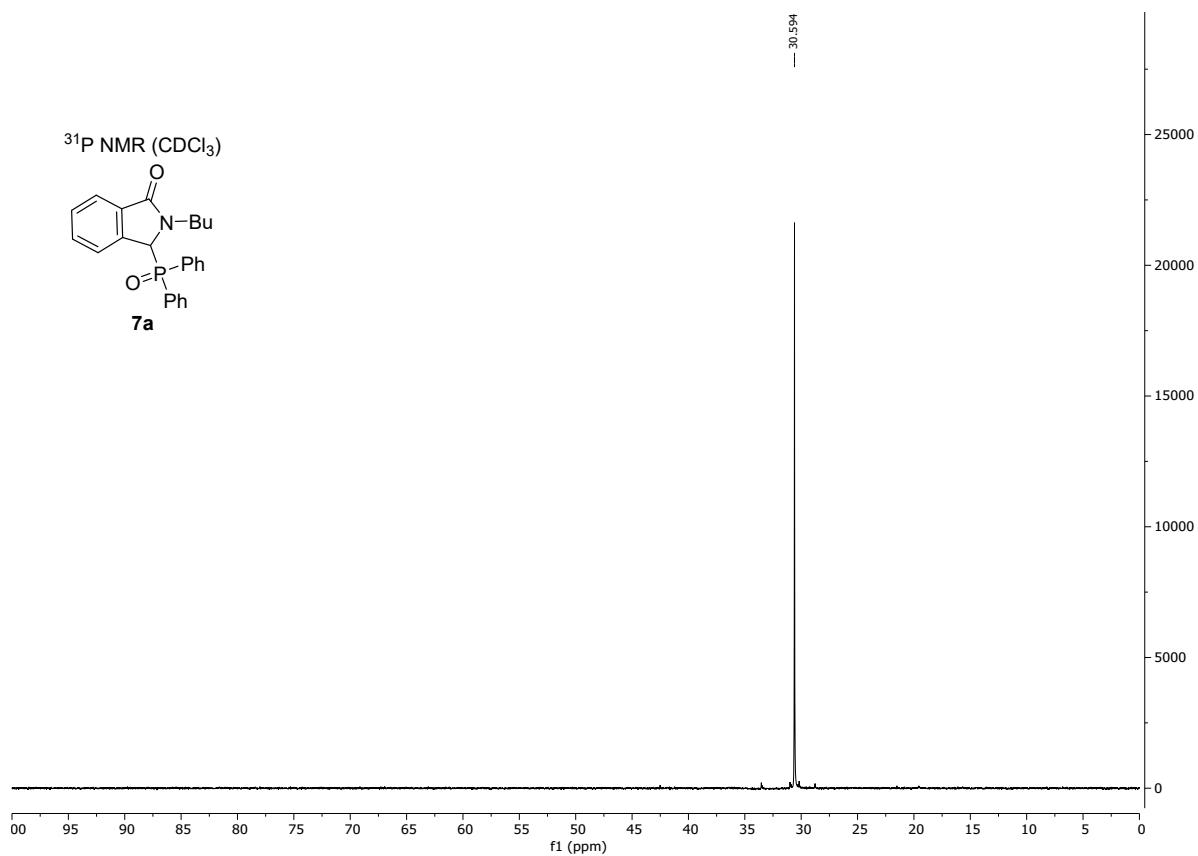
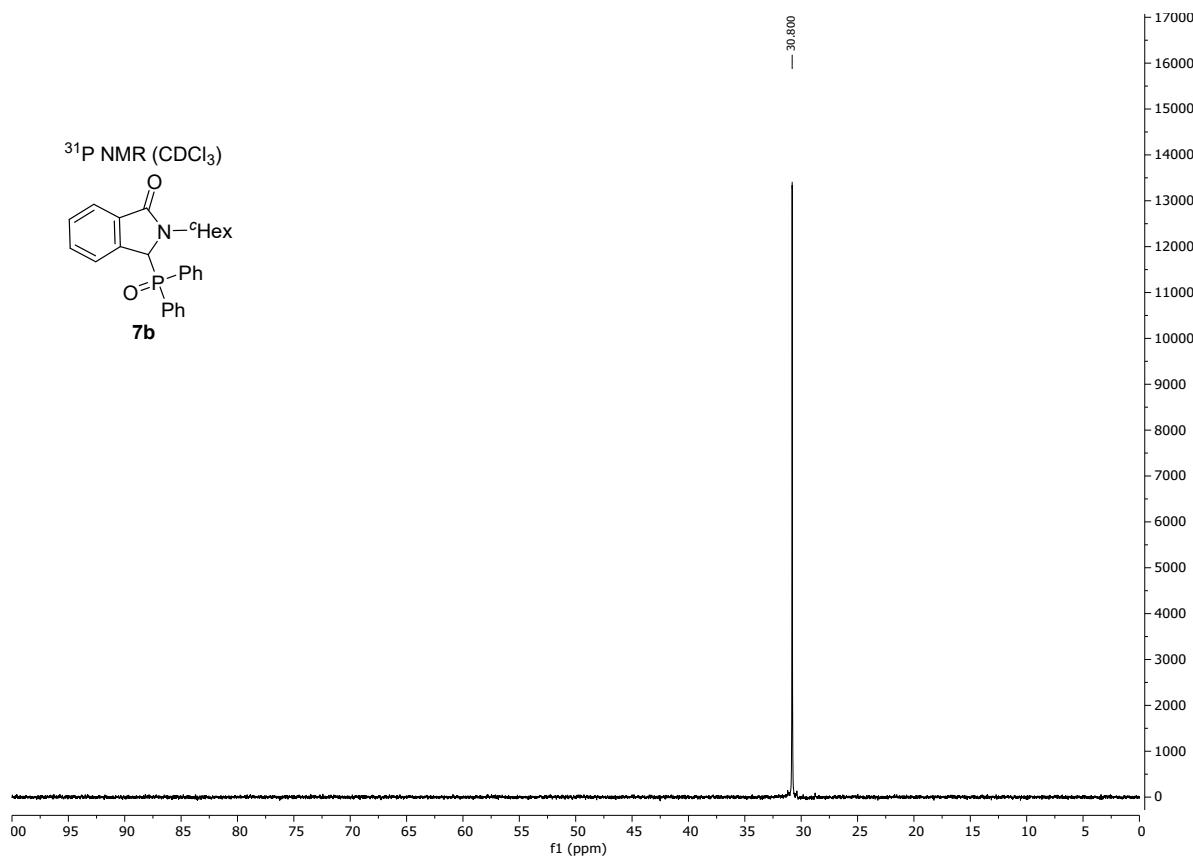
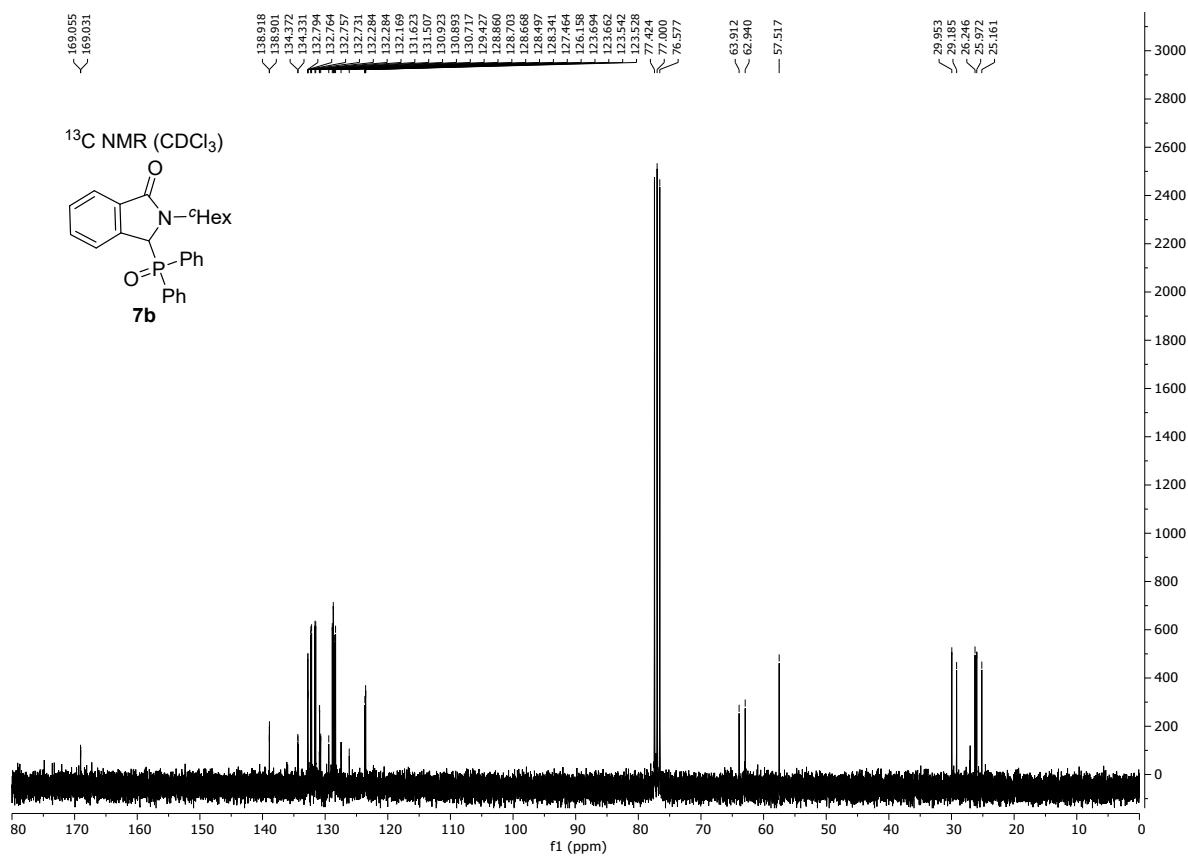


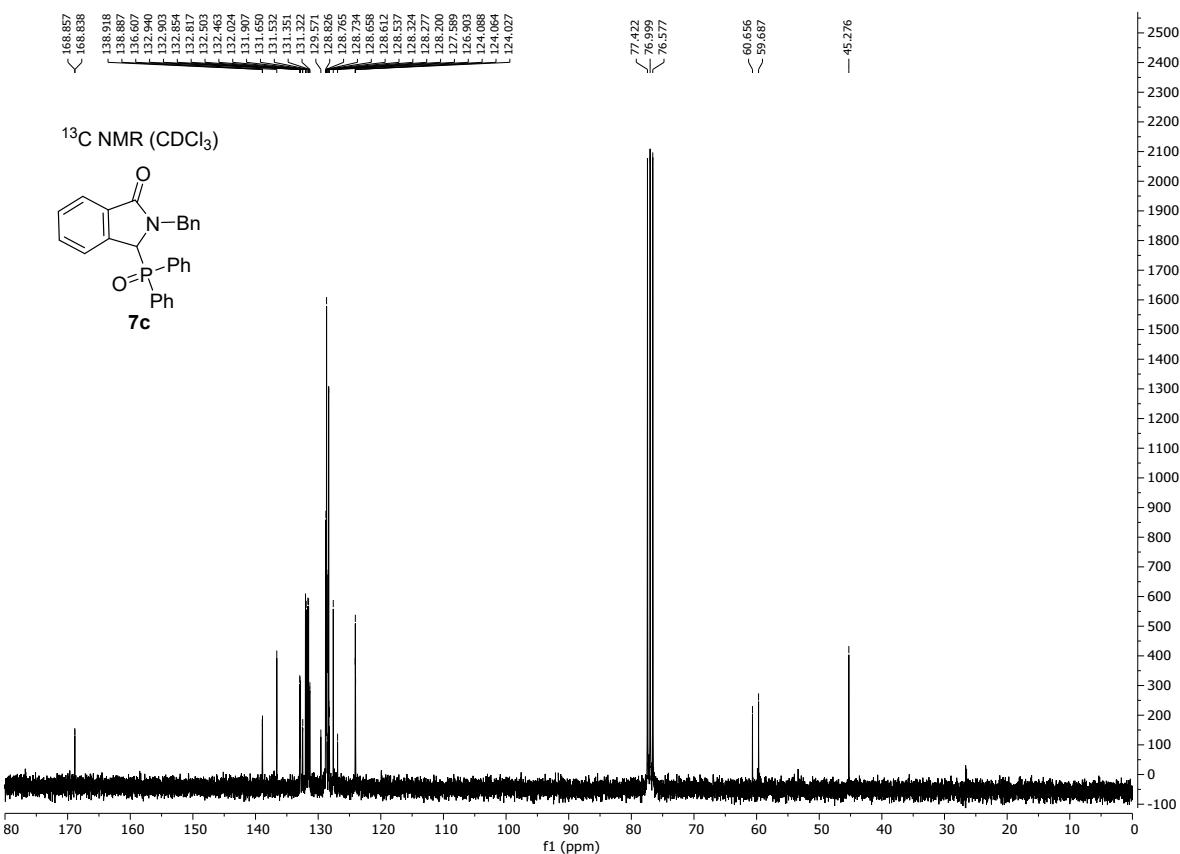
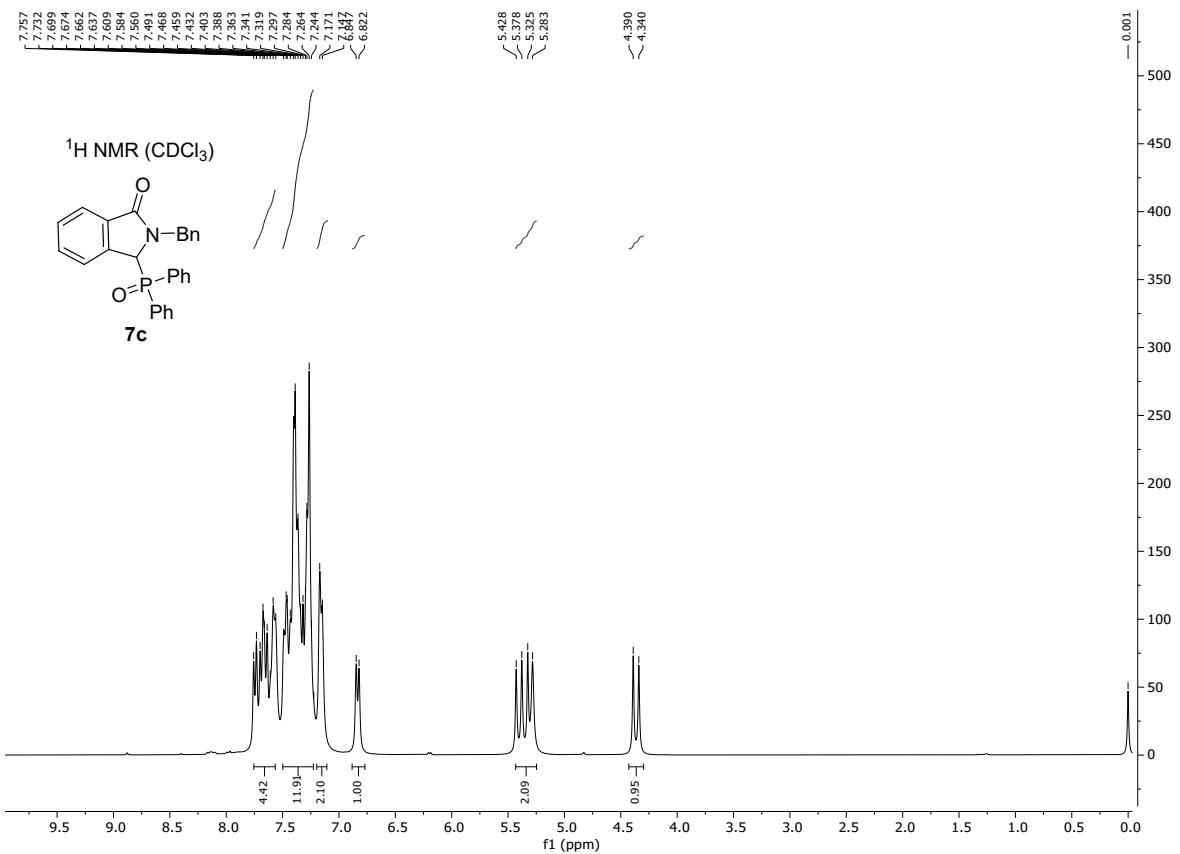
Figure S2. (a) Hydrogen-bonded layer formation in **22** via C21–H21···O1 interactions. (b) C24–H24···Cg2 interactions. (c) Packing of layers (arbitrary colours). Blue dashed lines indicate hydrogen bonds.

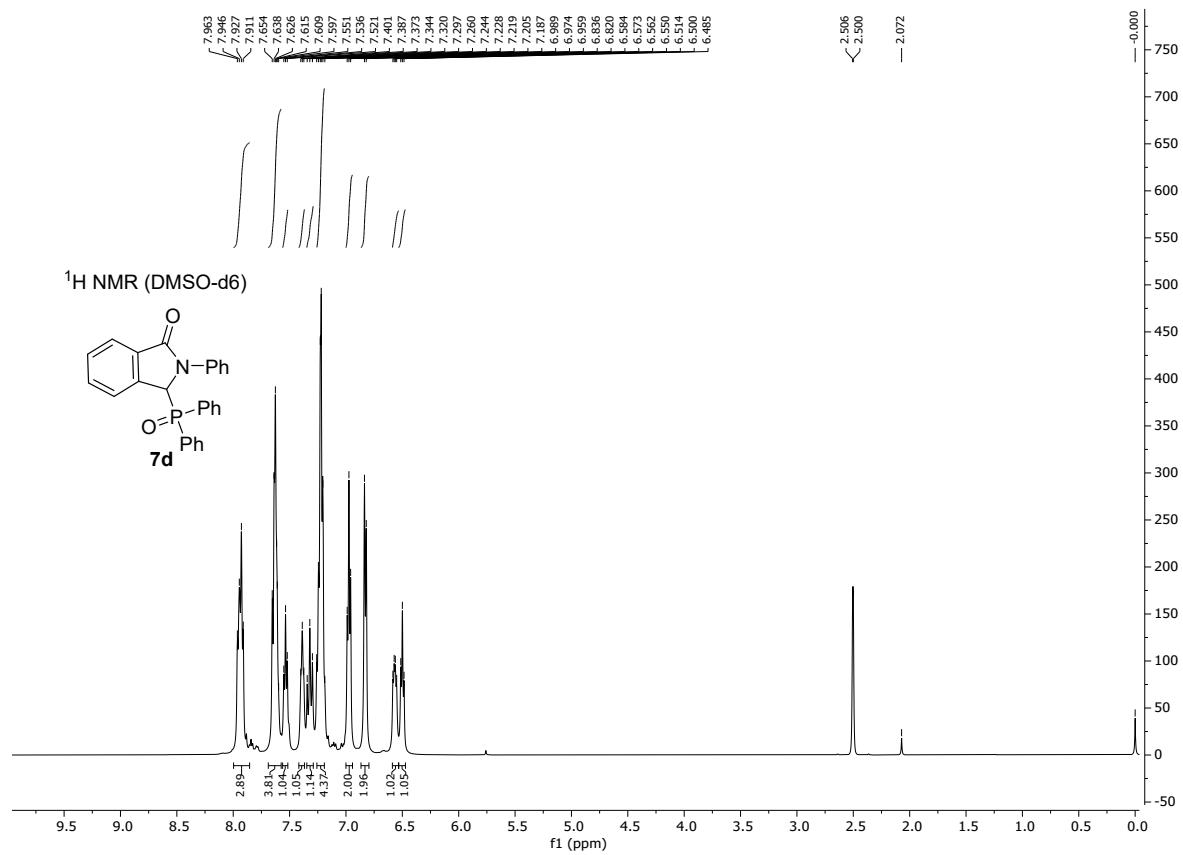
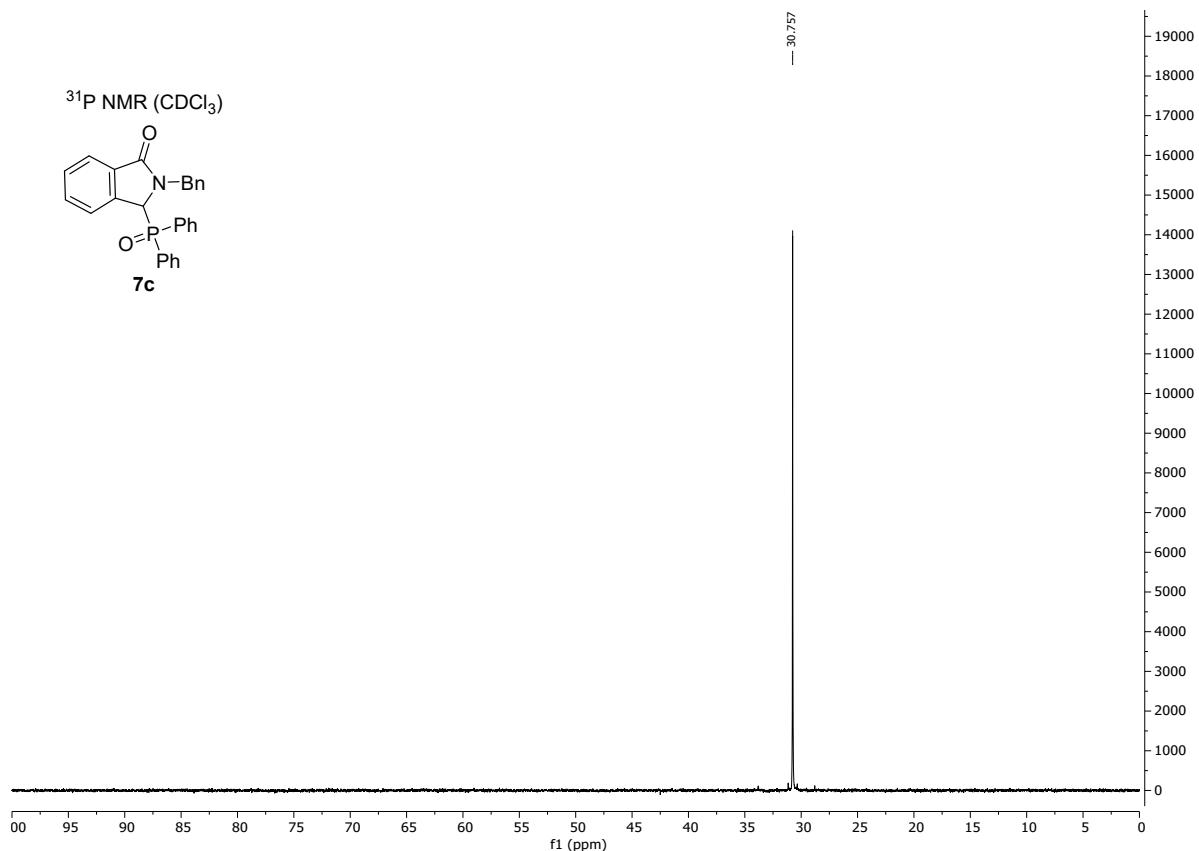
¹H NMR, ¹³C NMR and ³¹P NMR spectra

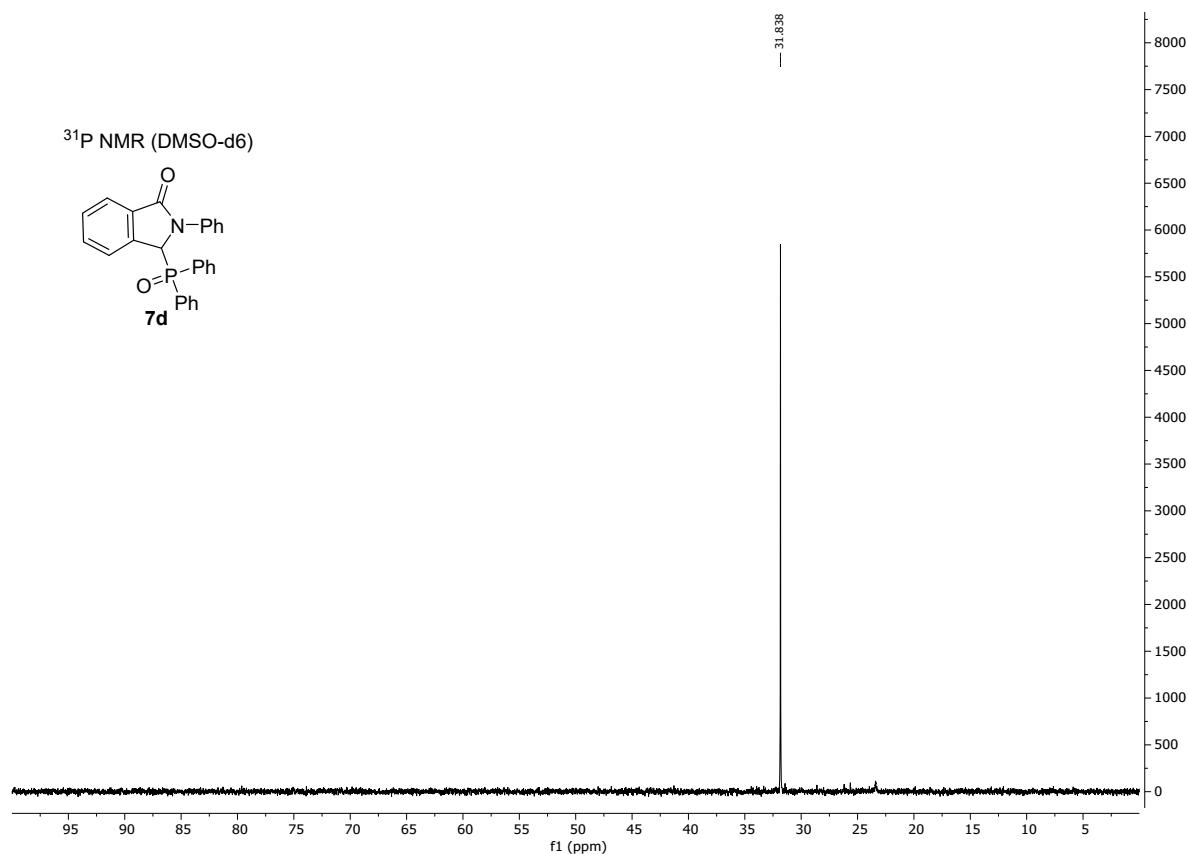
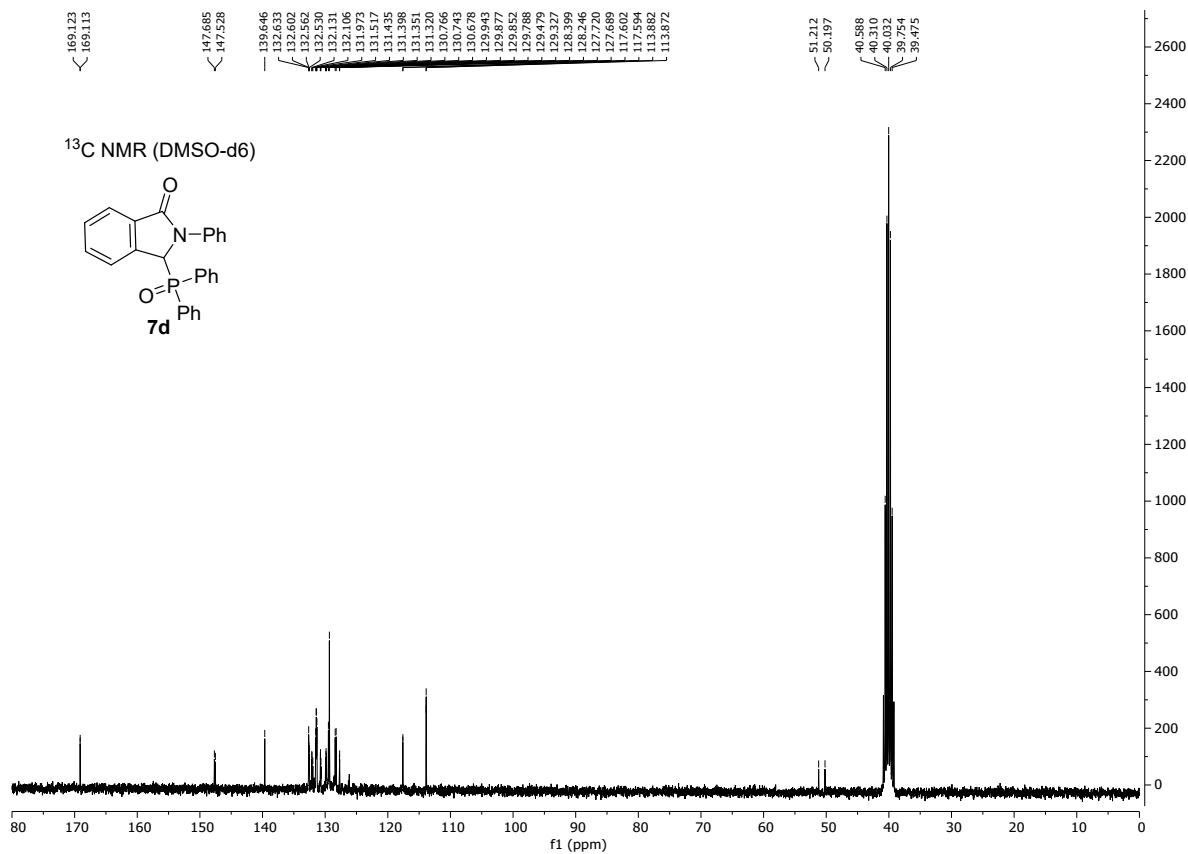


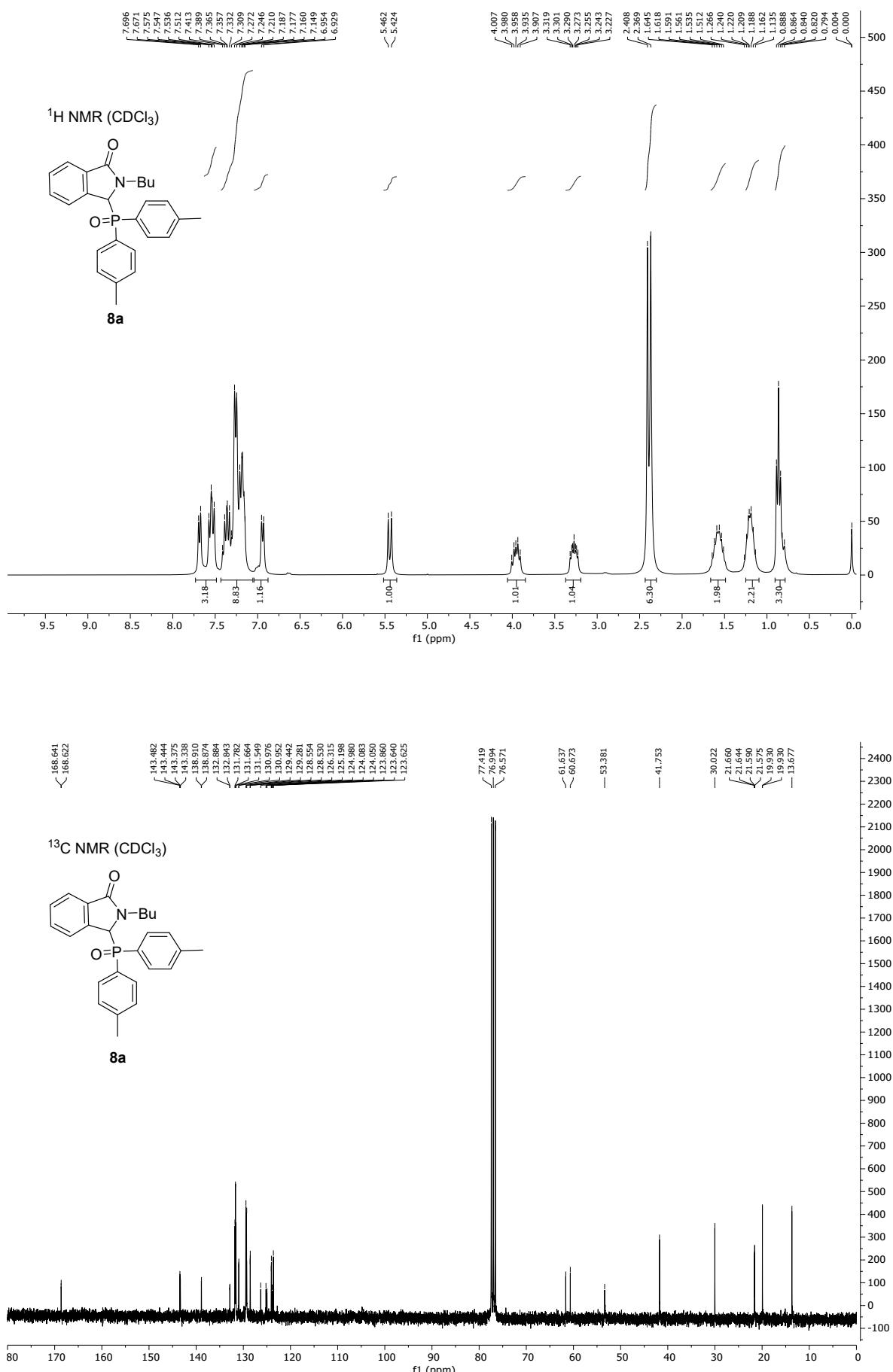


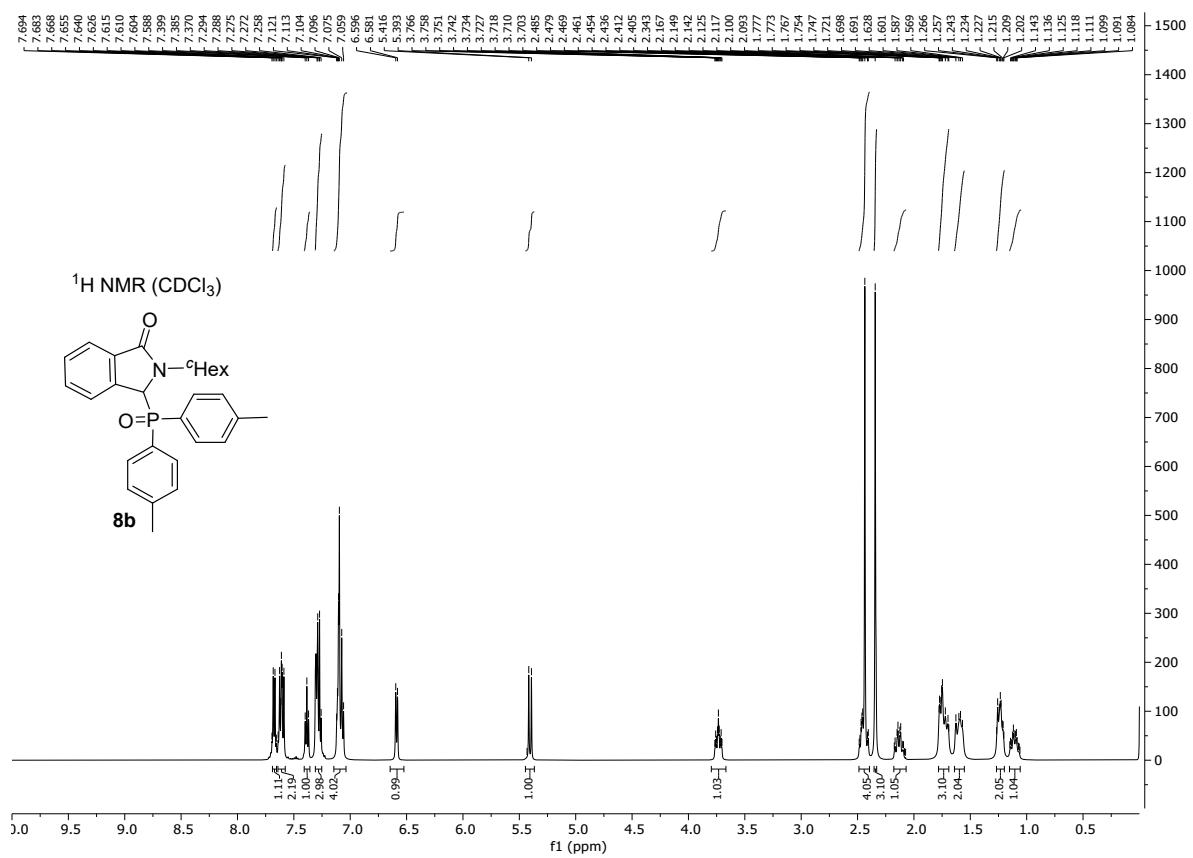
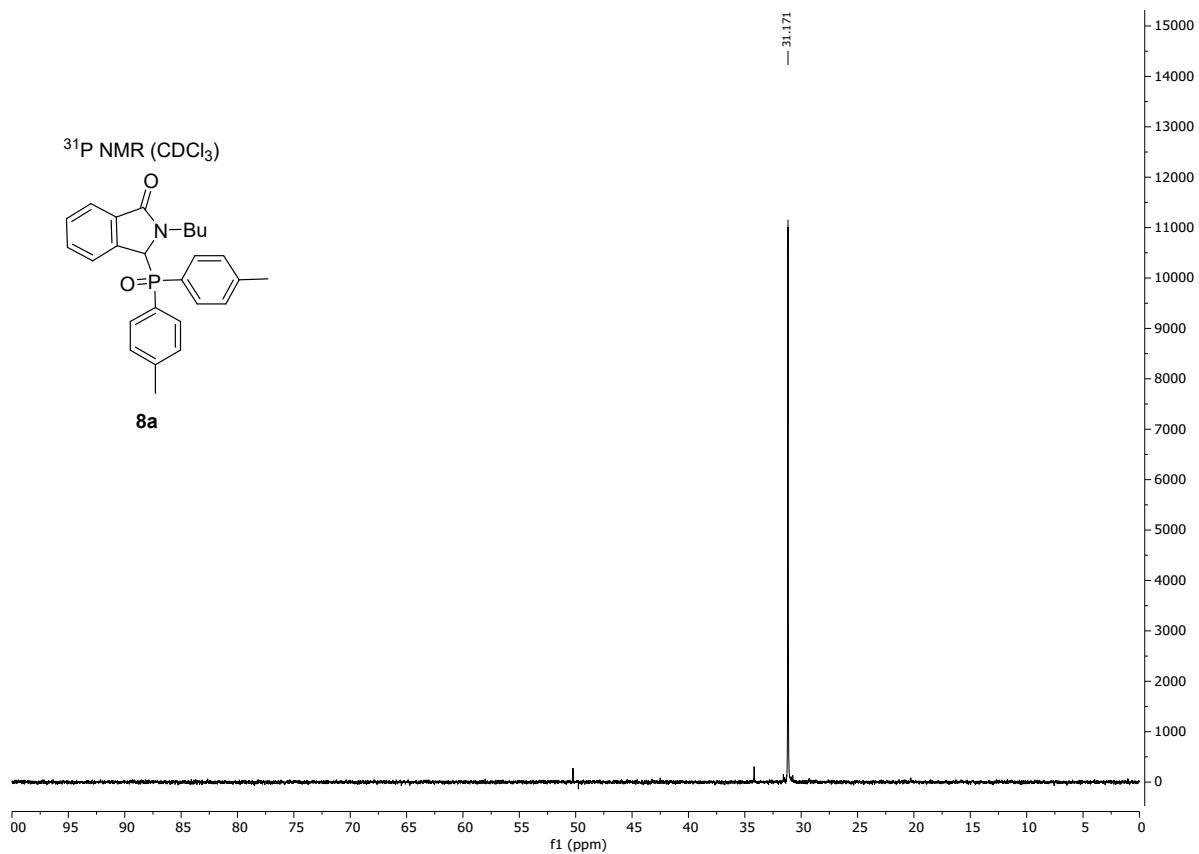


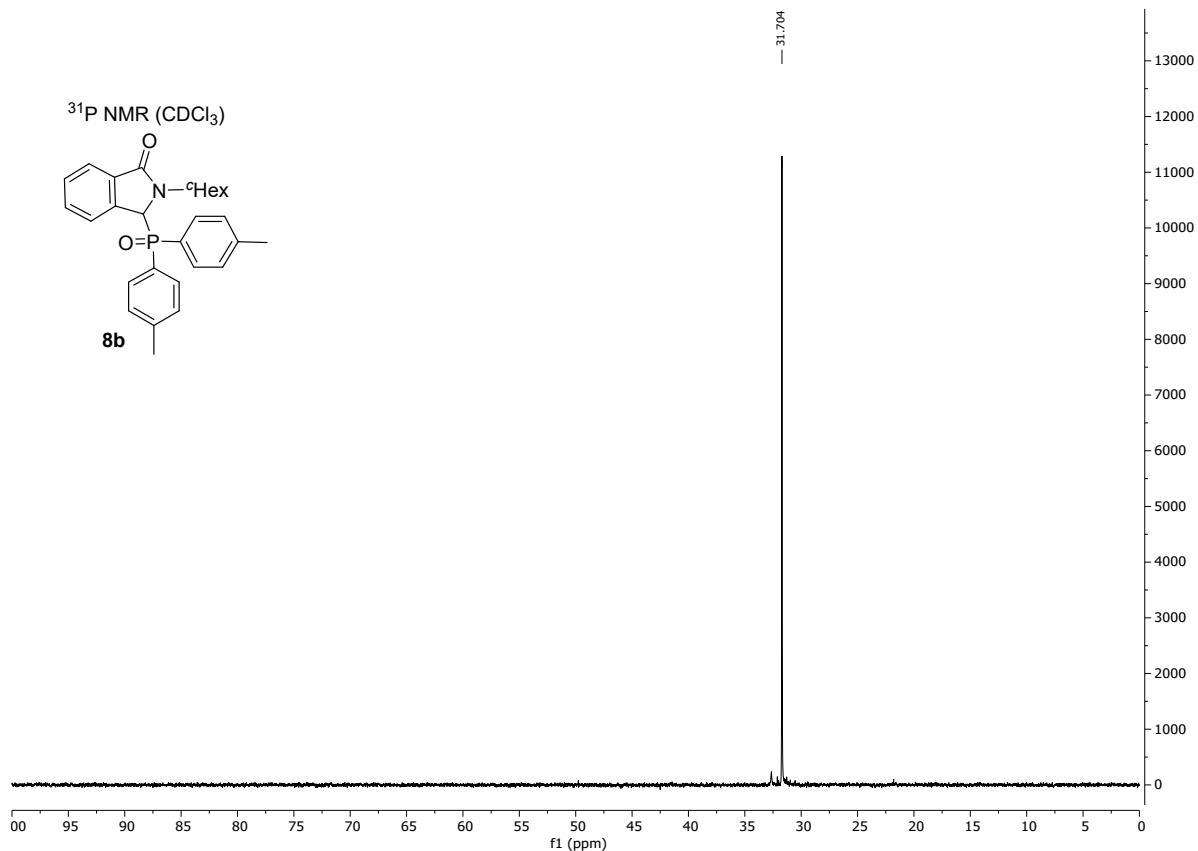
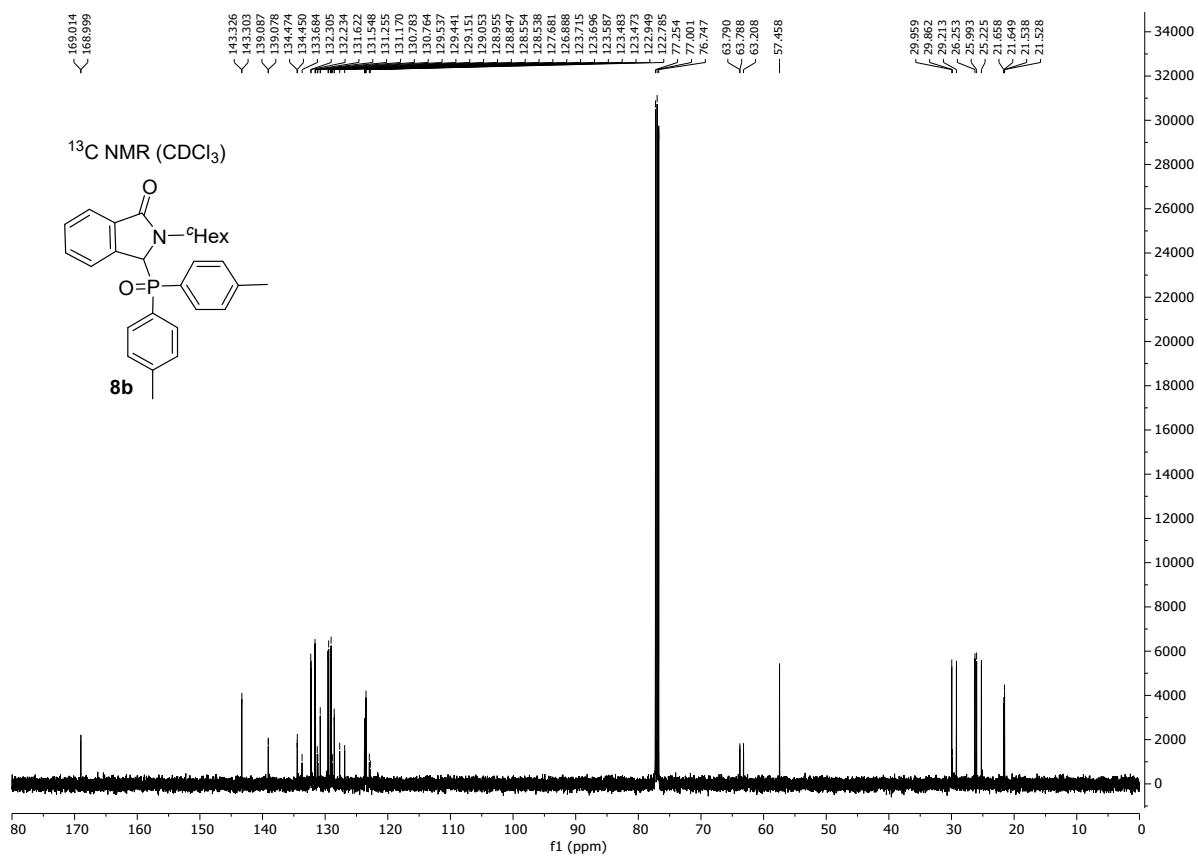


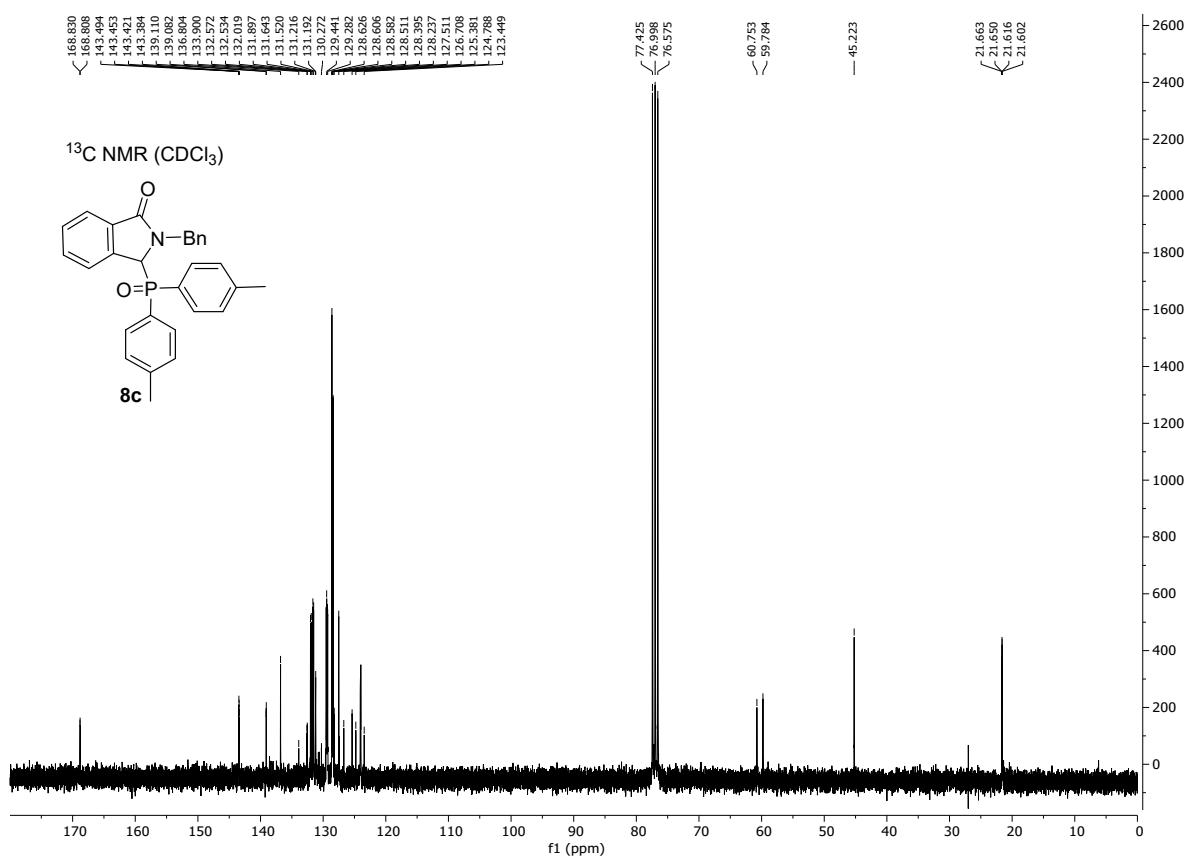
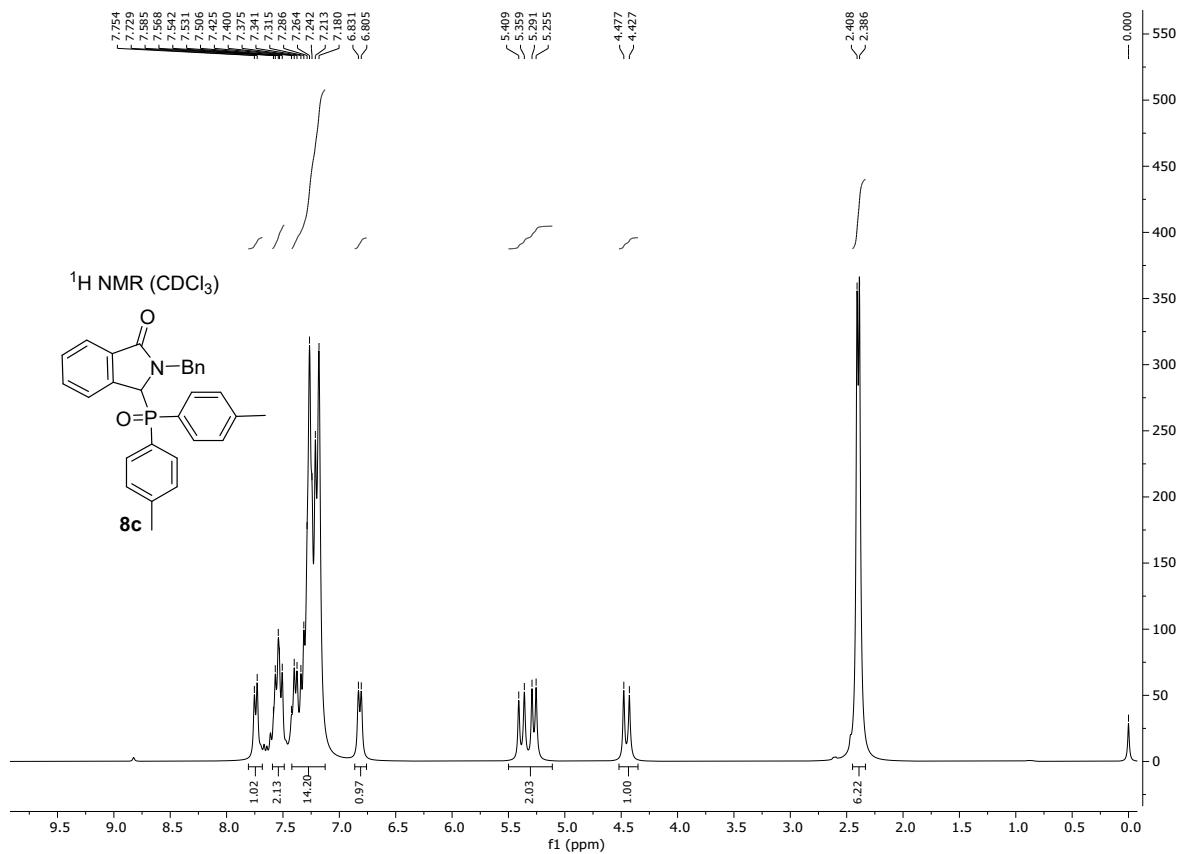


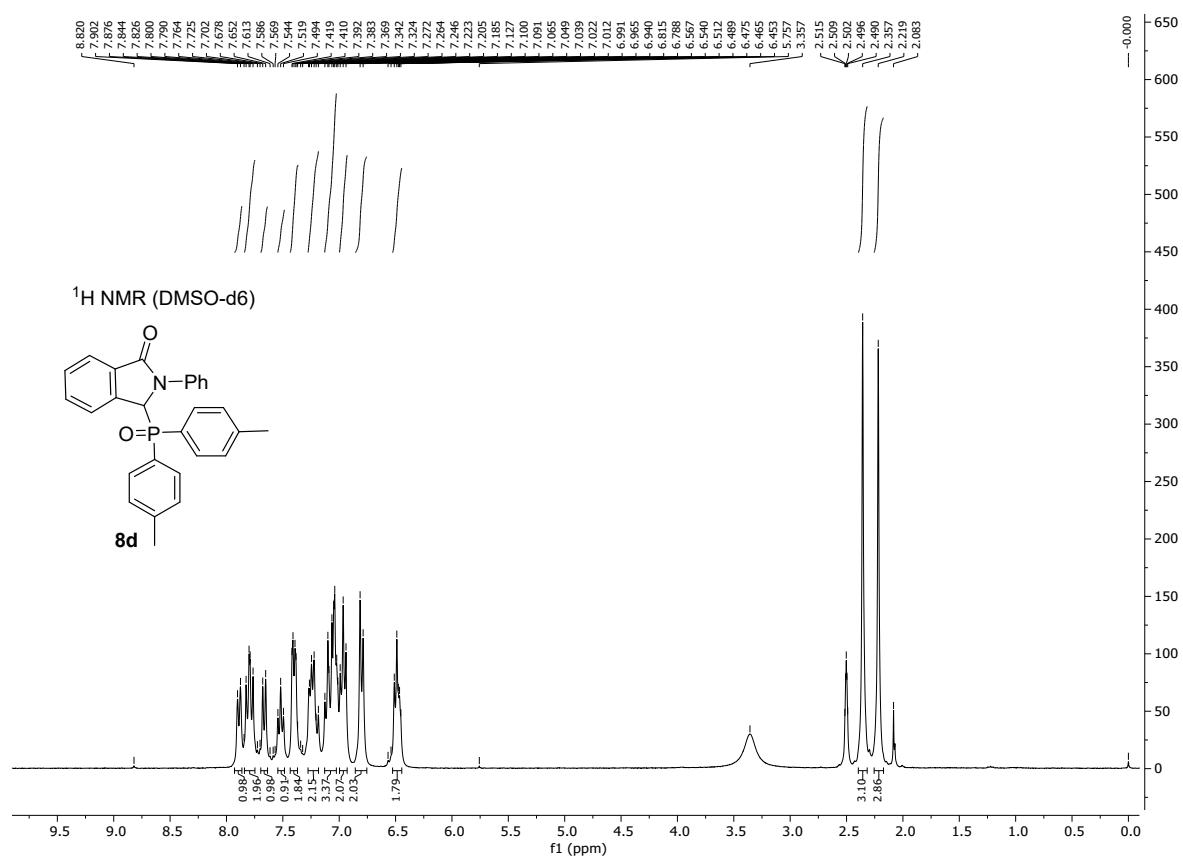
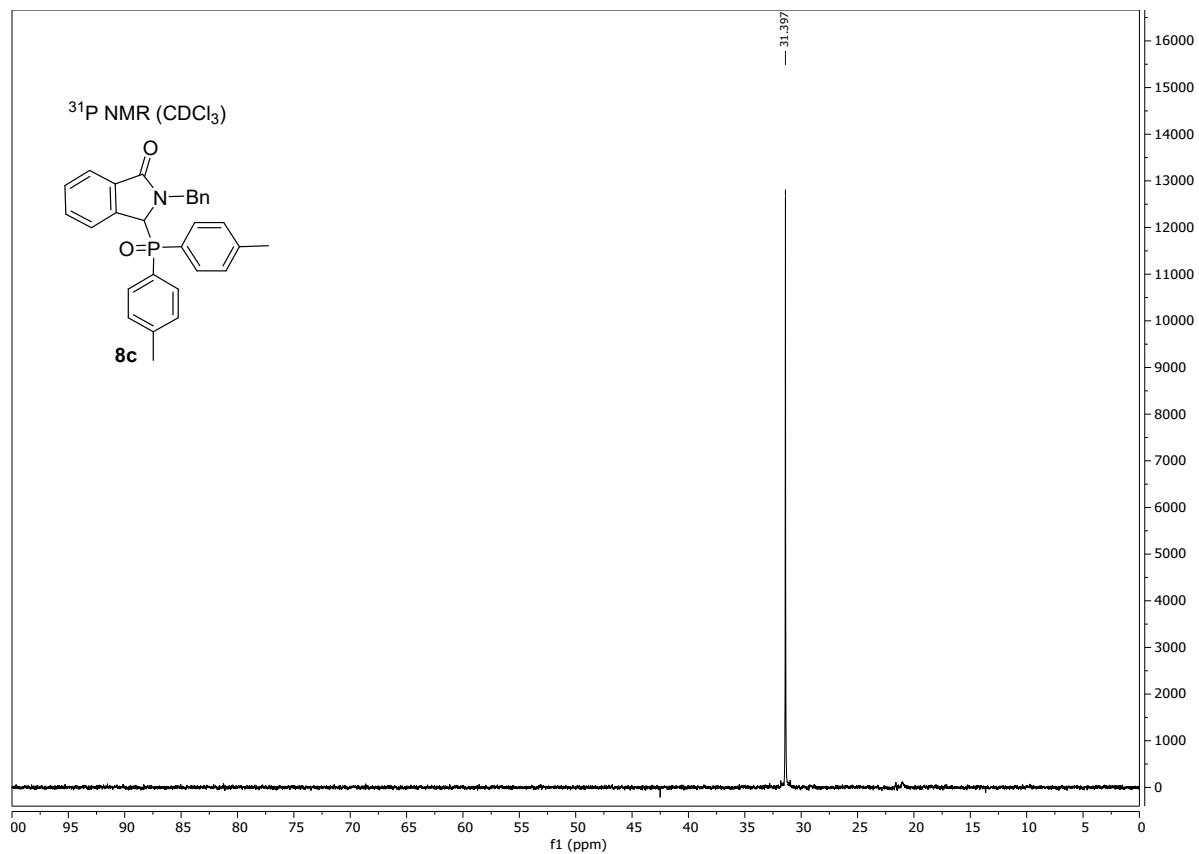


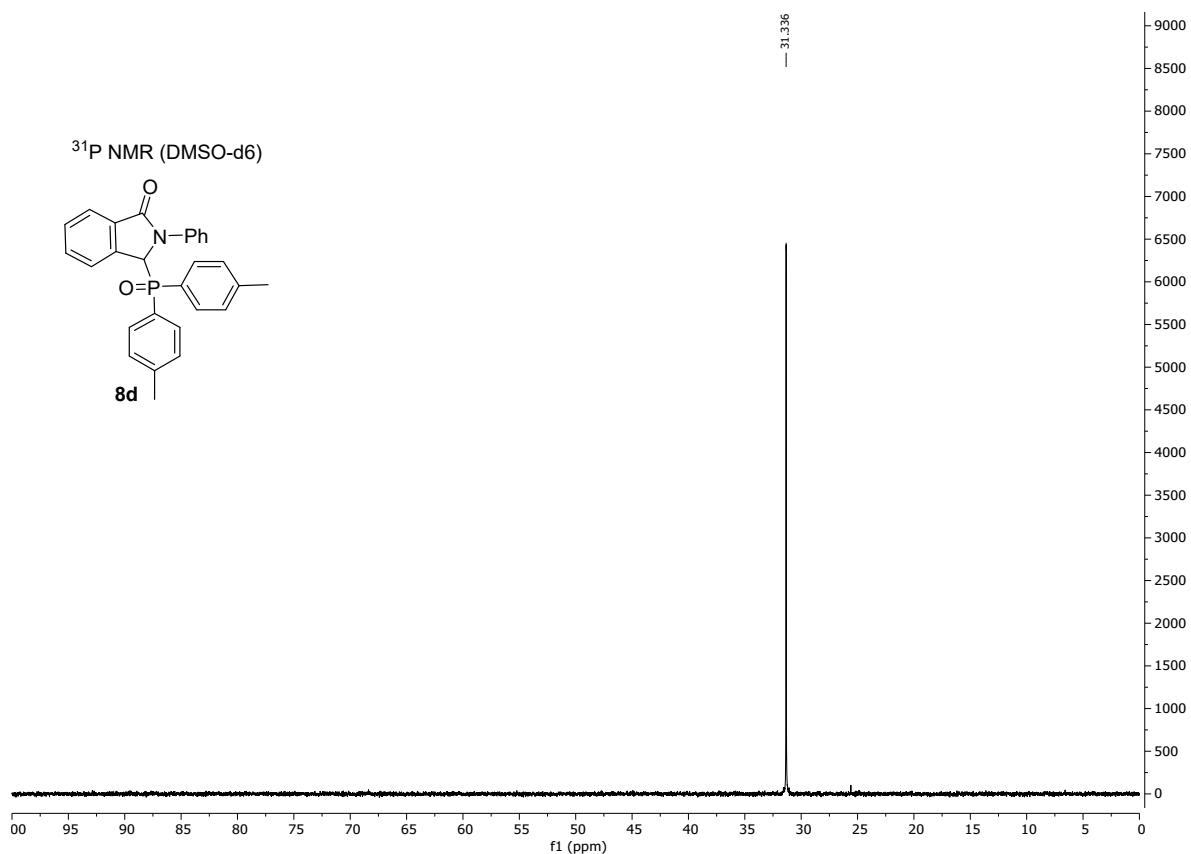
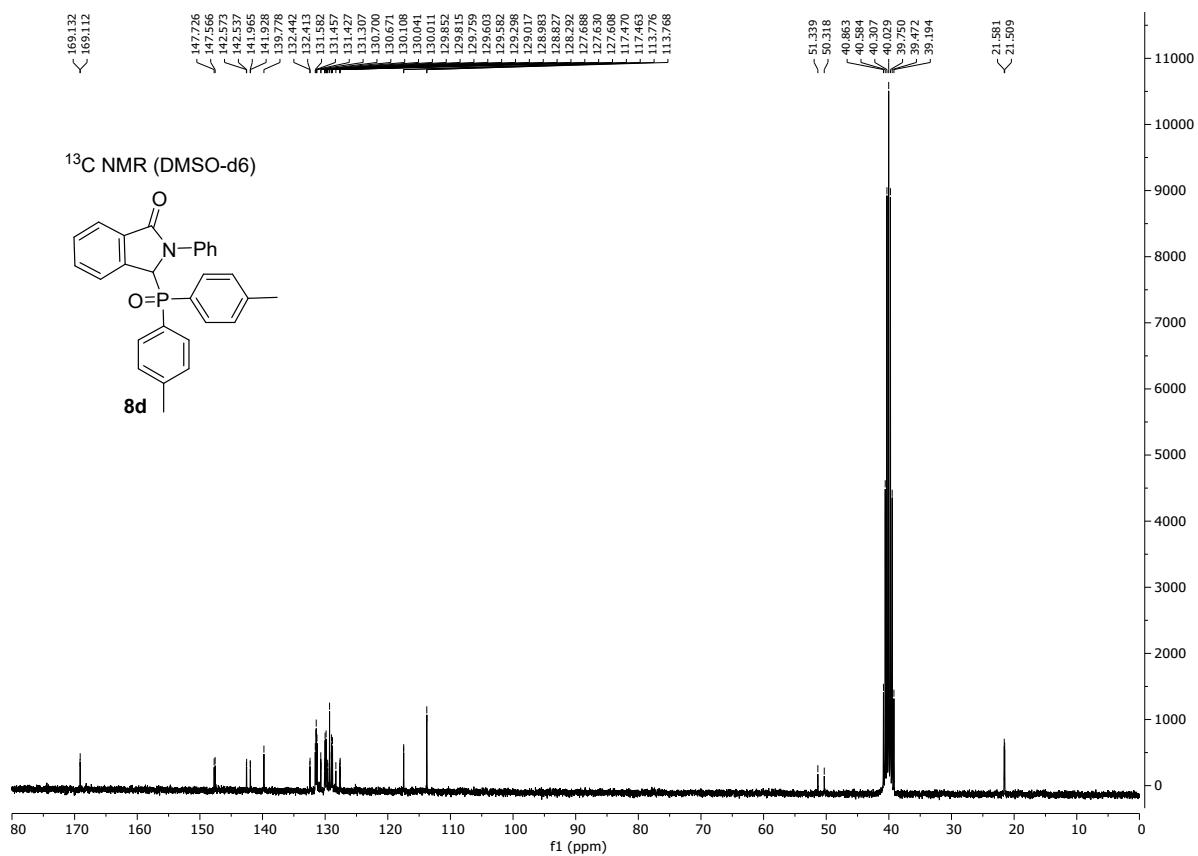


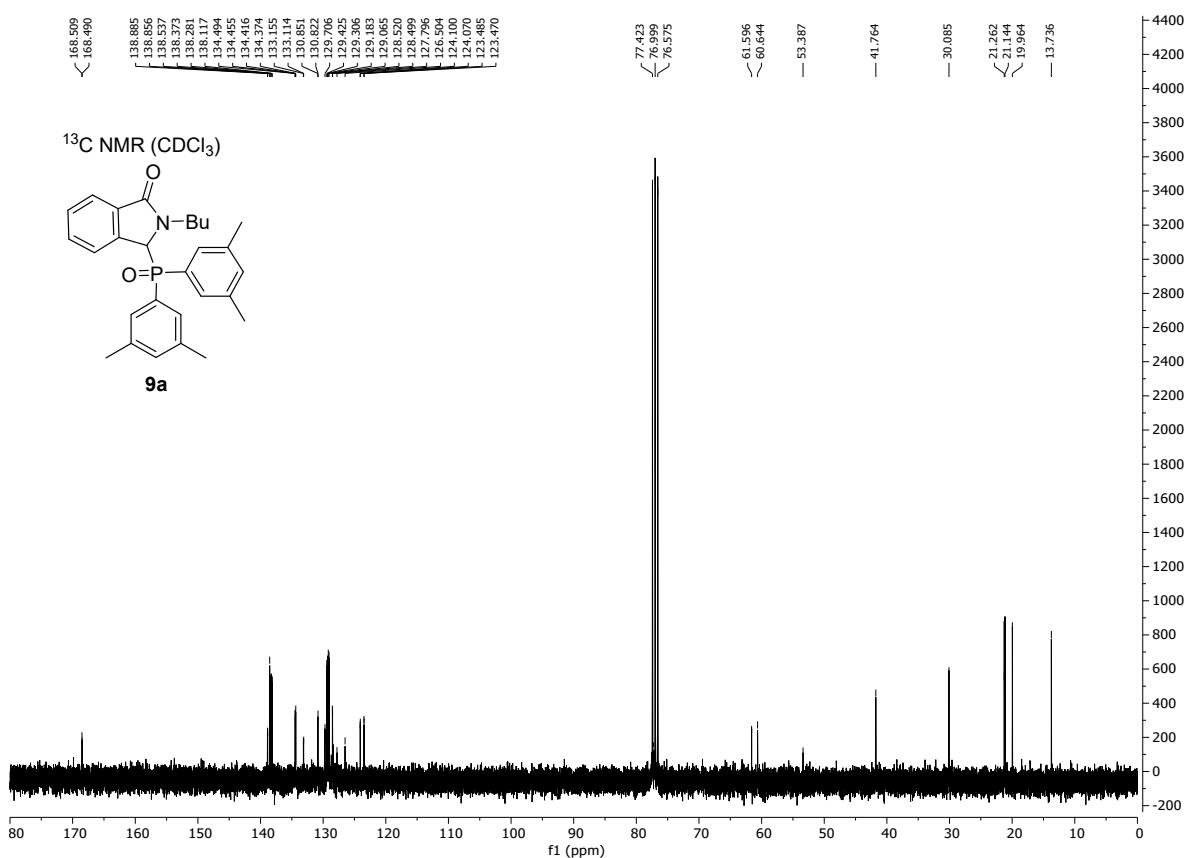
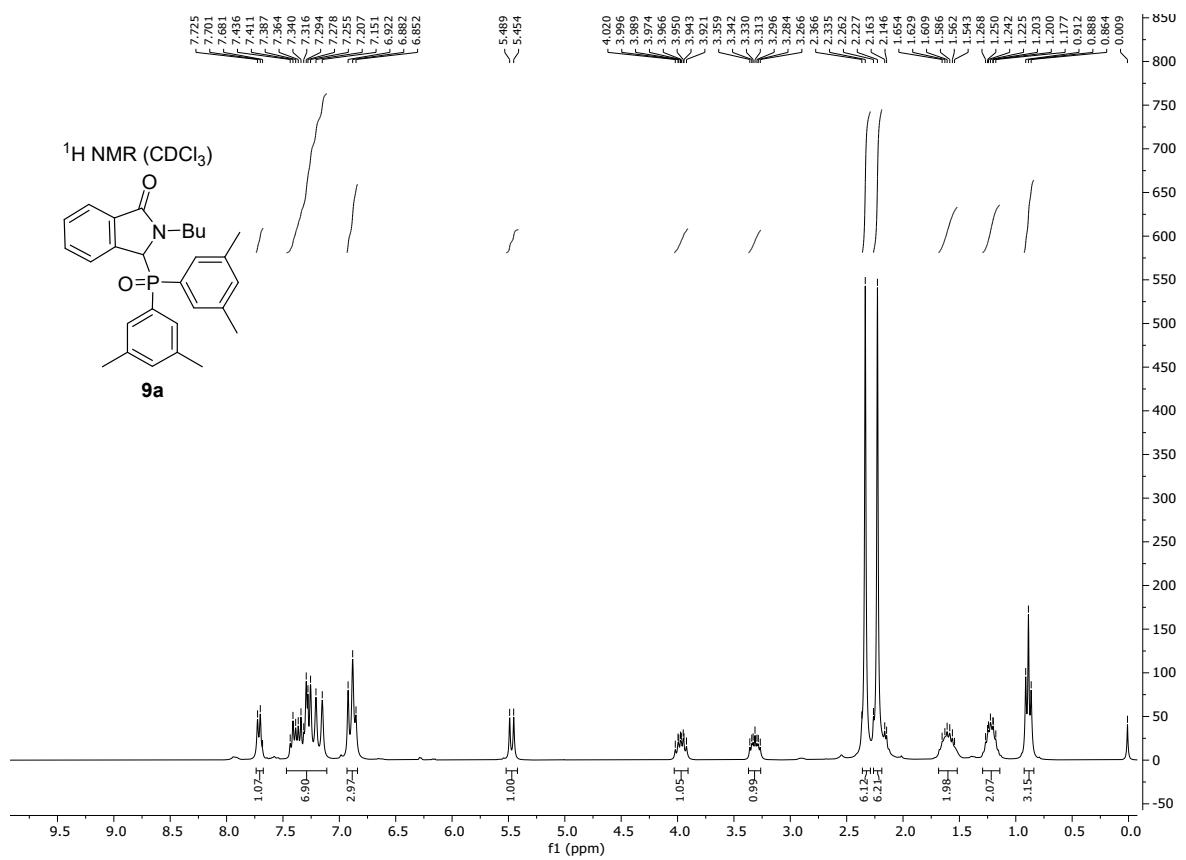


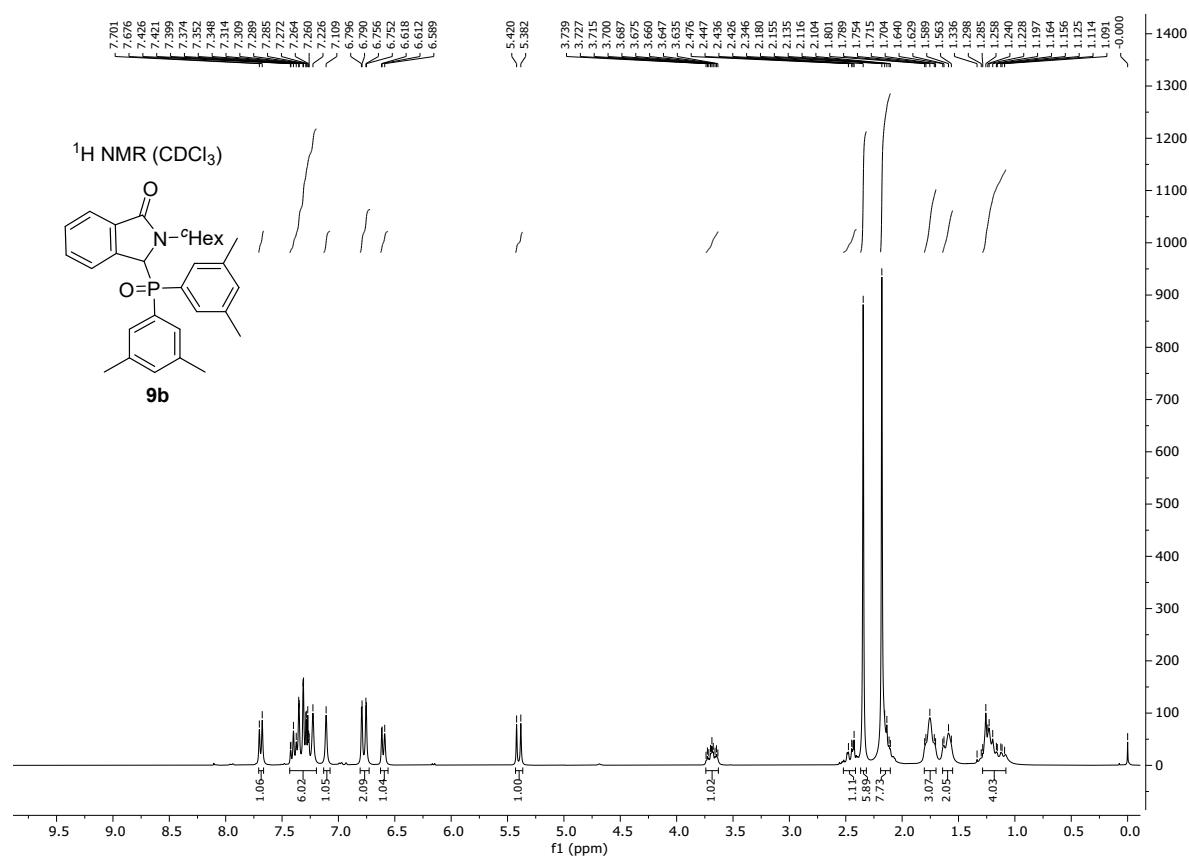
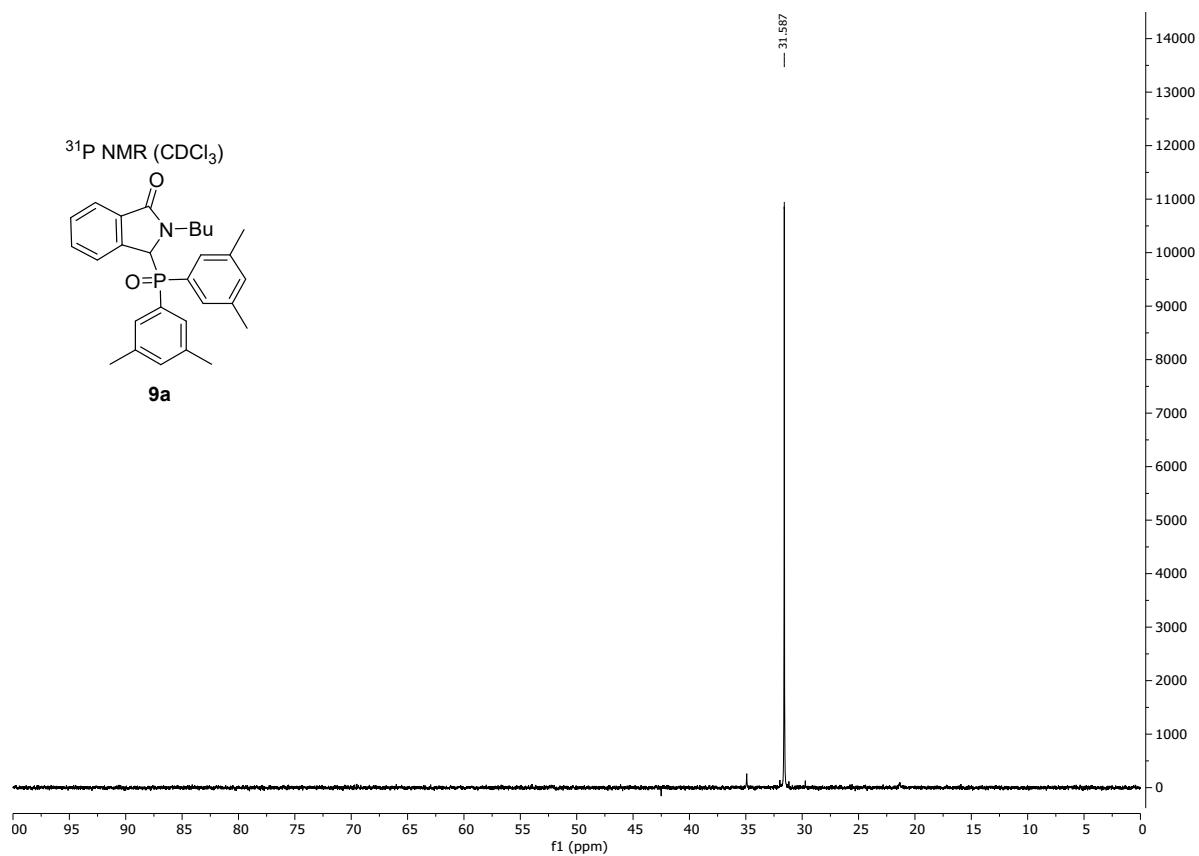


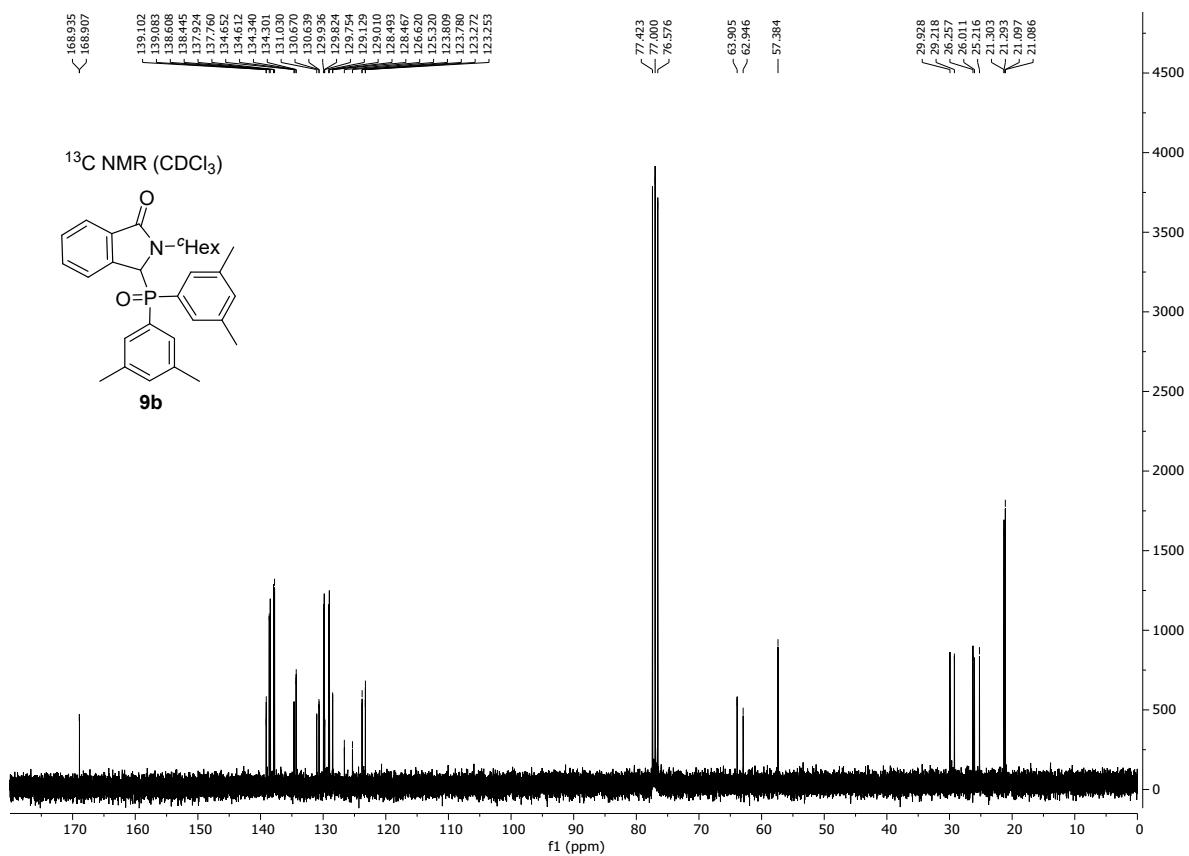


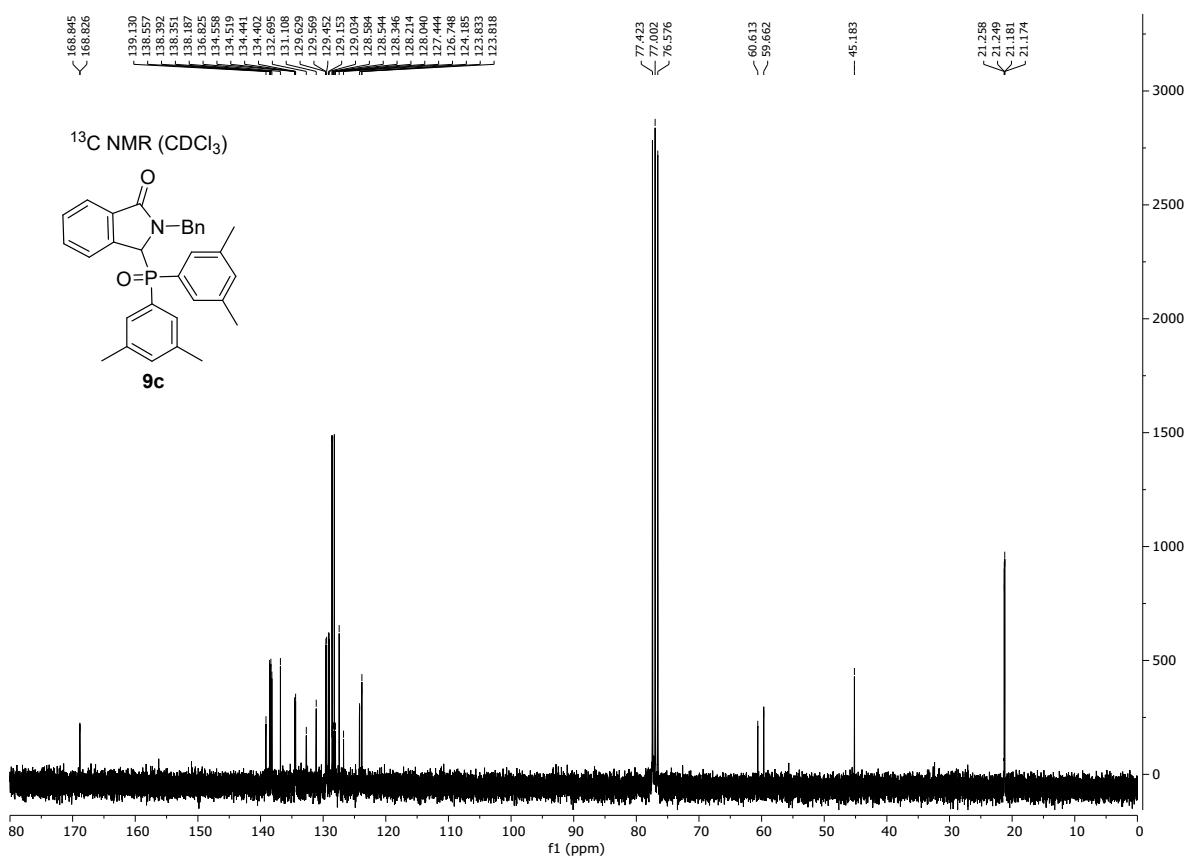
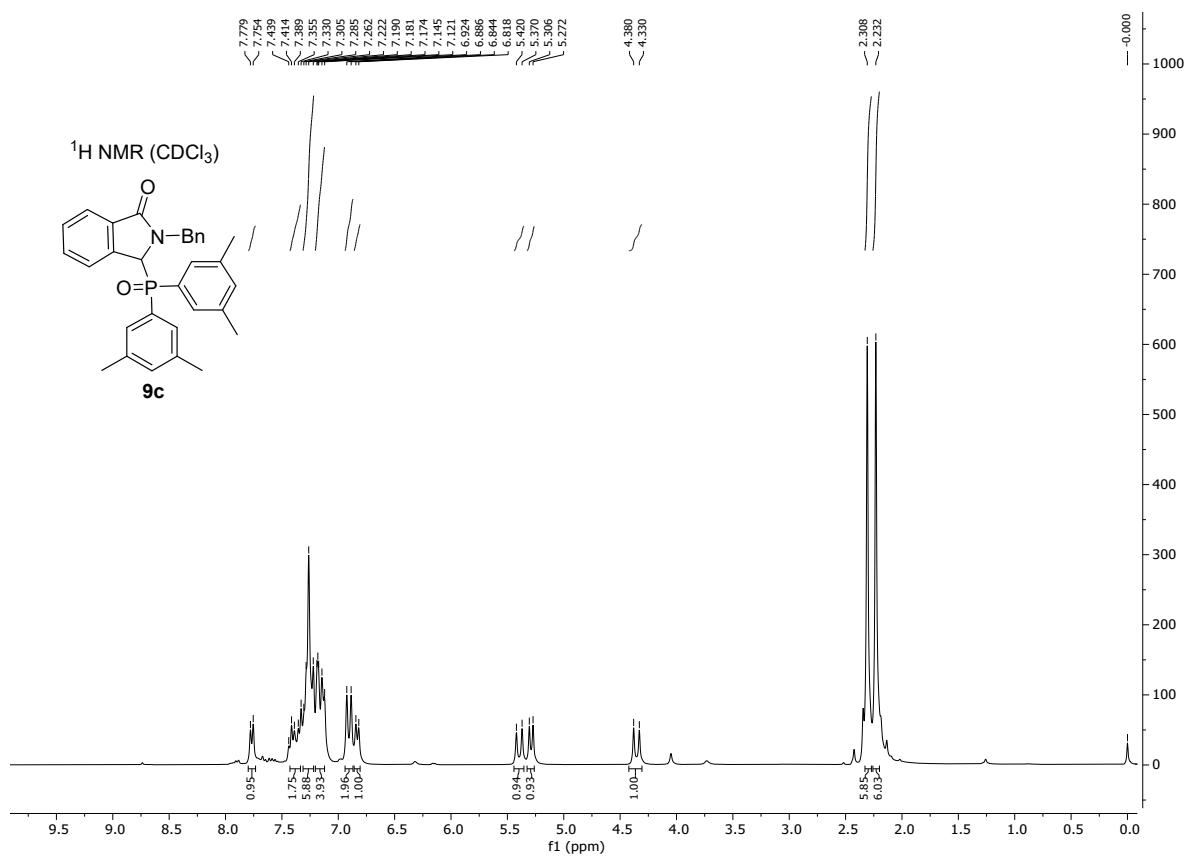


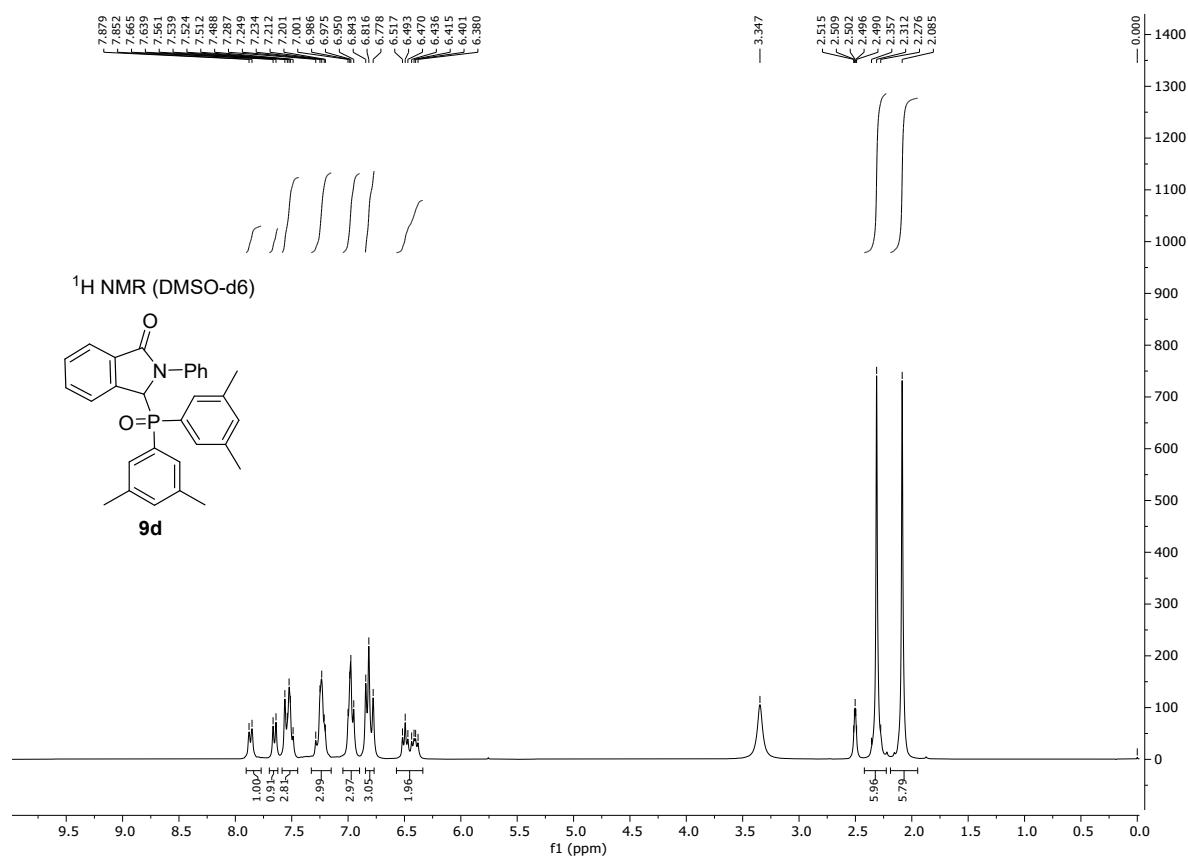
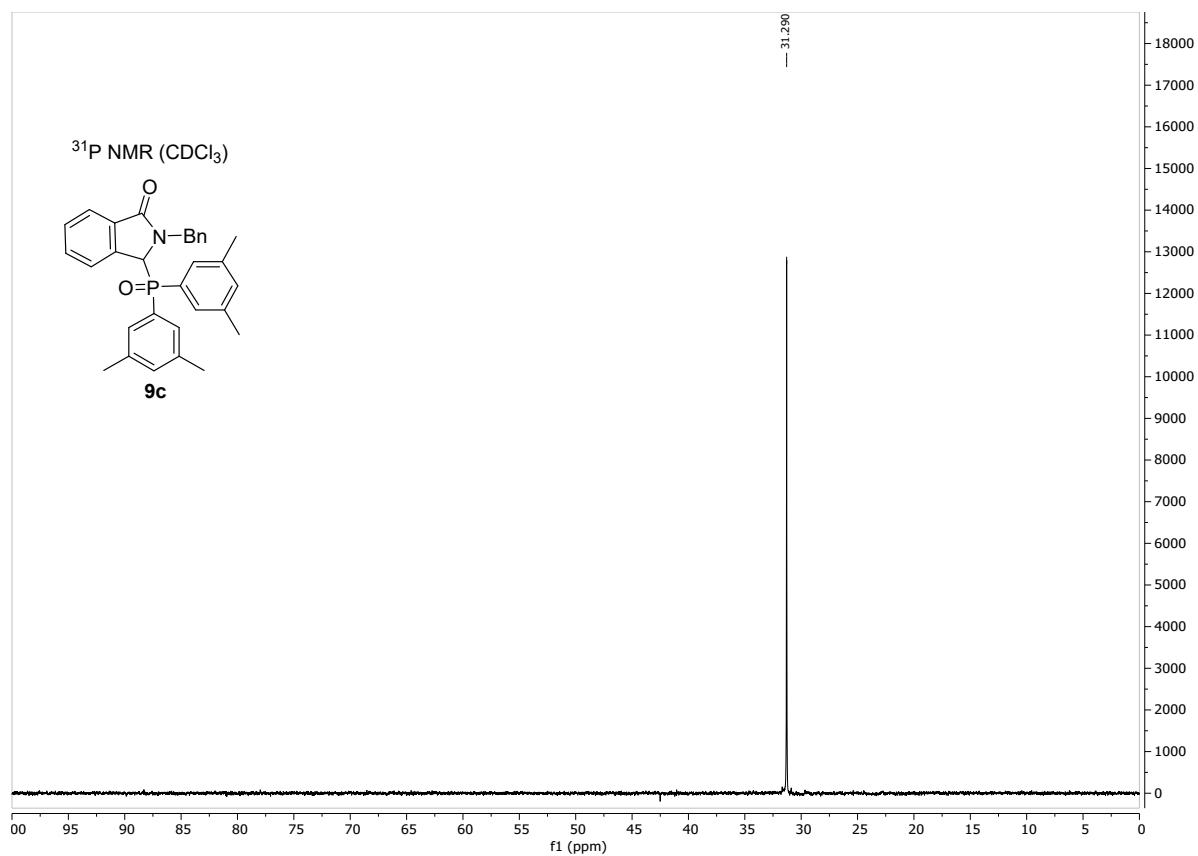


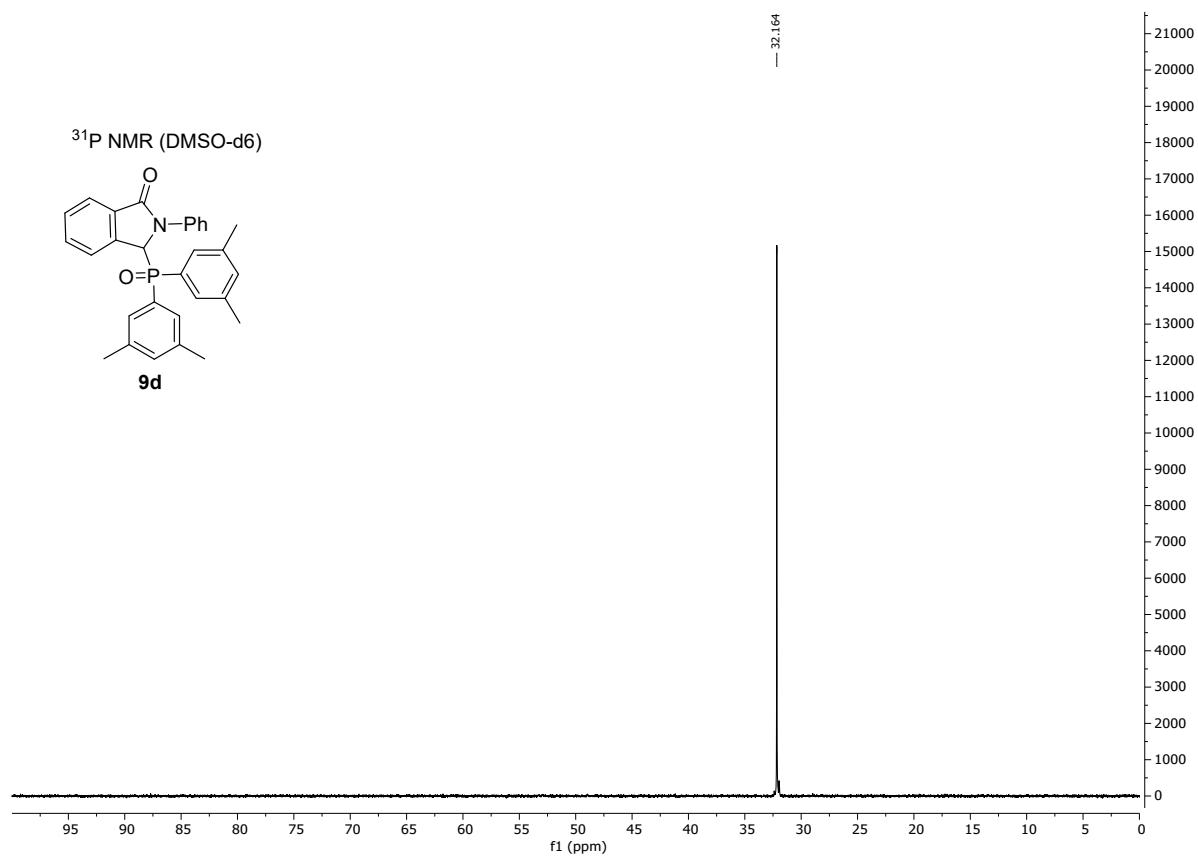
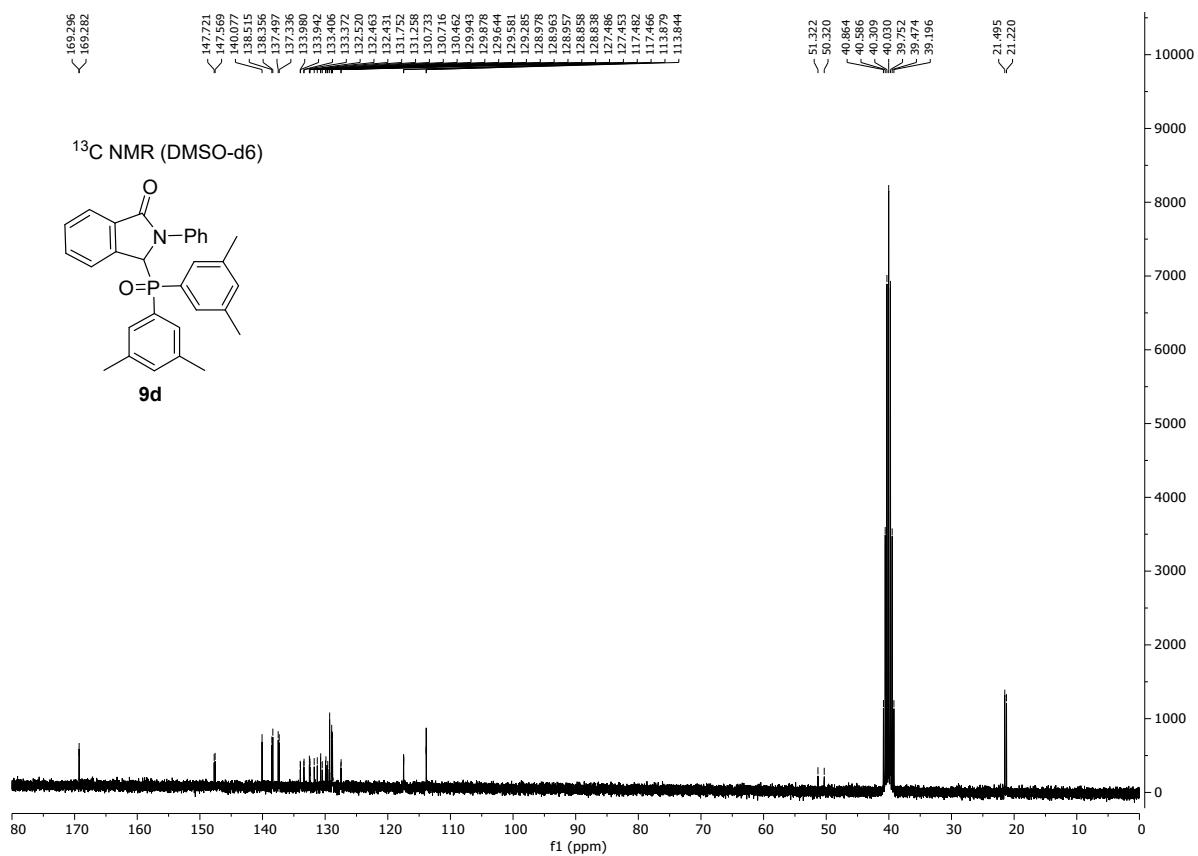


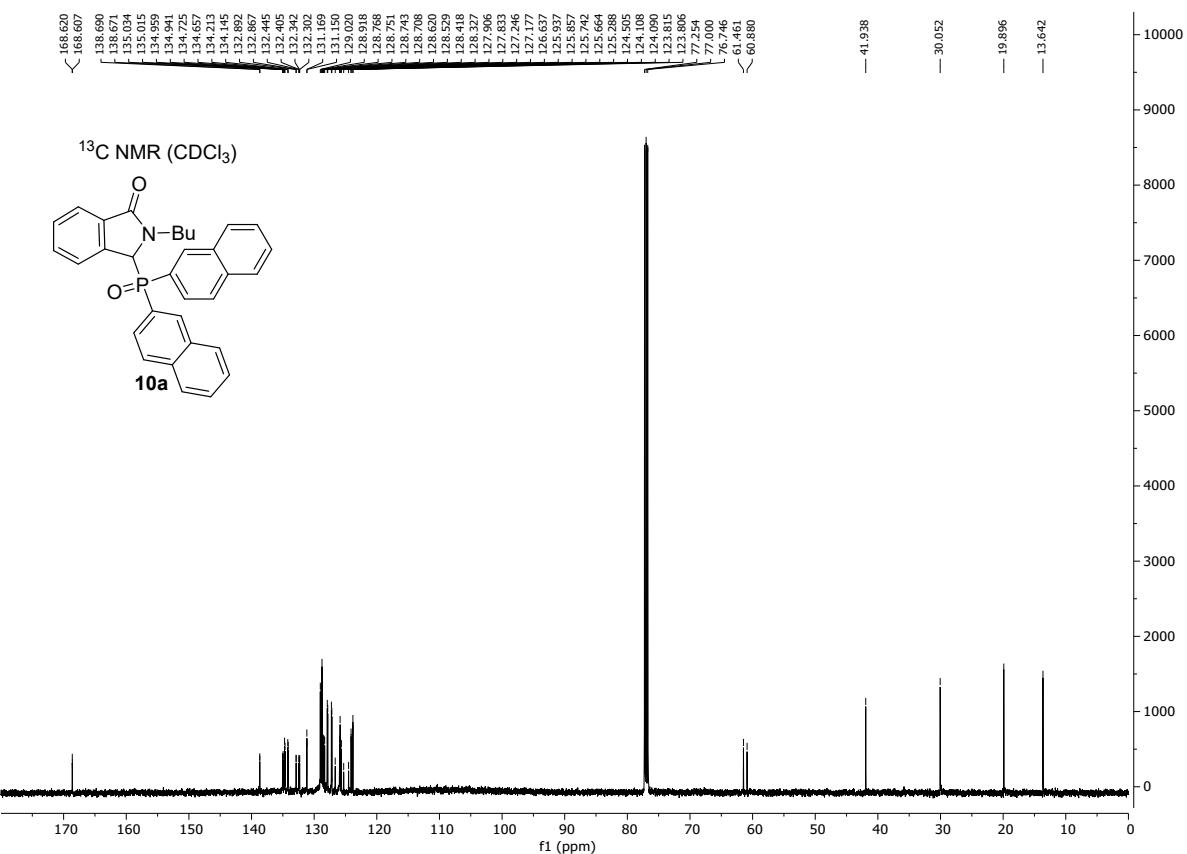
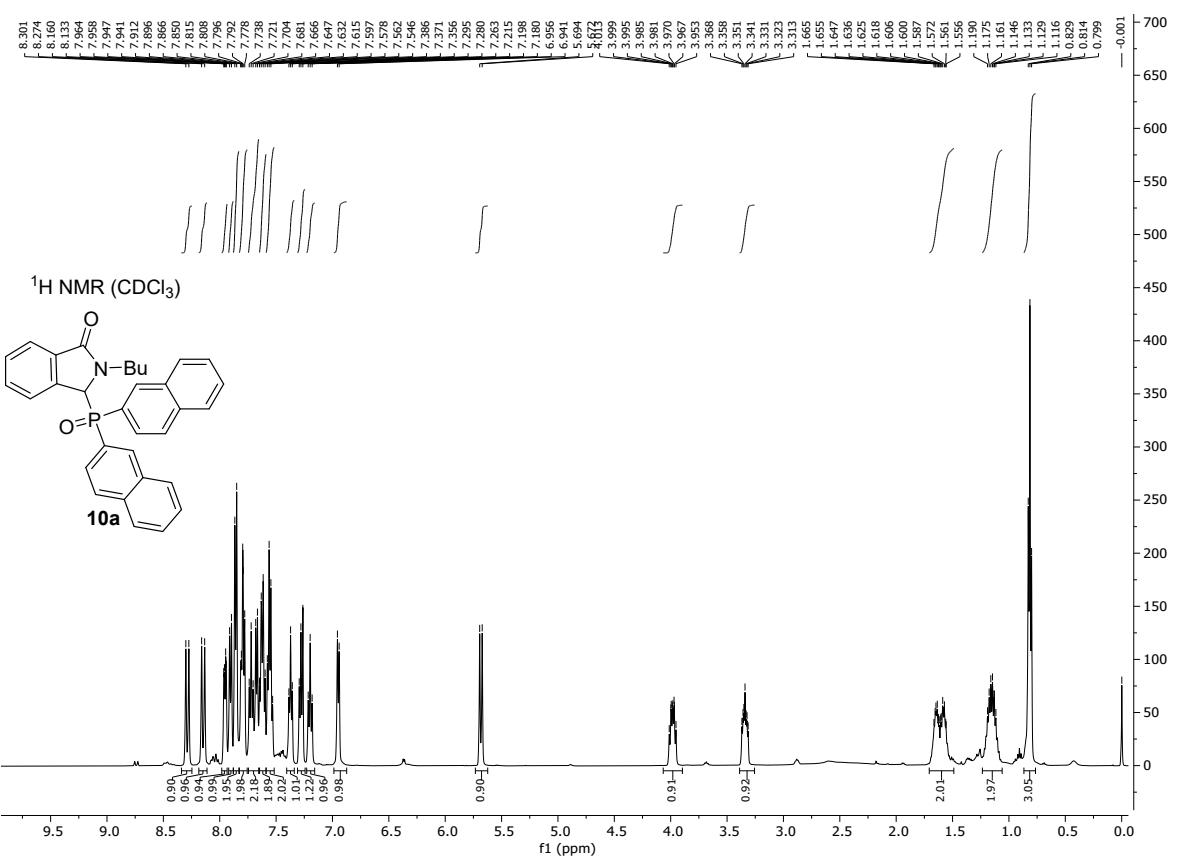


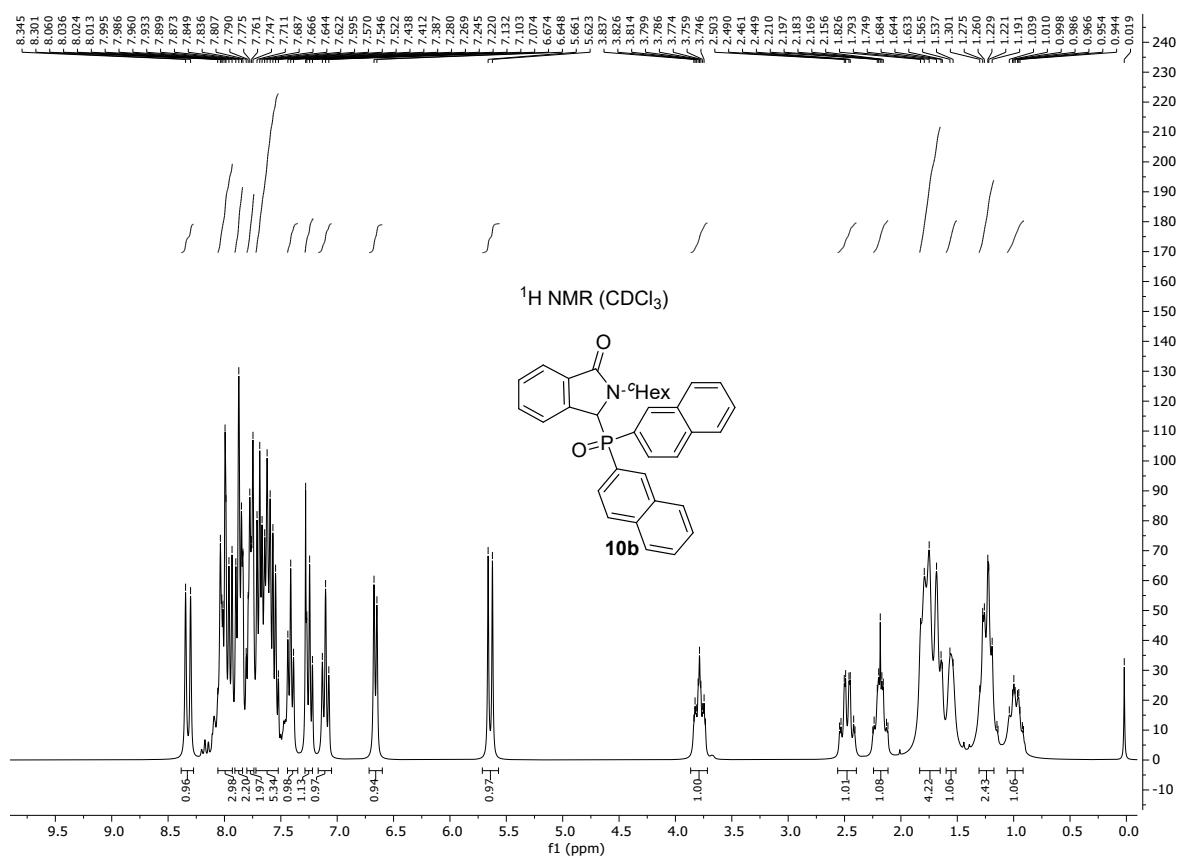
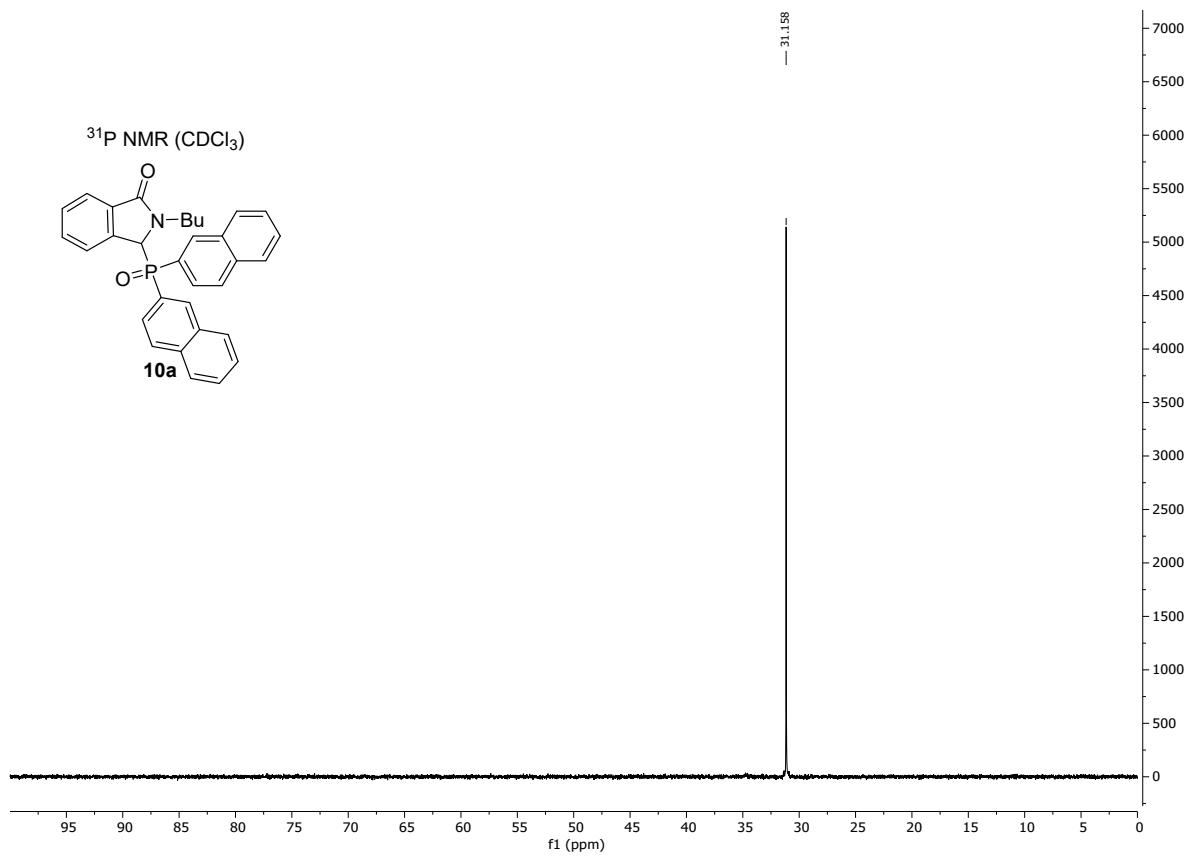


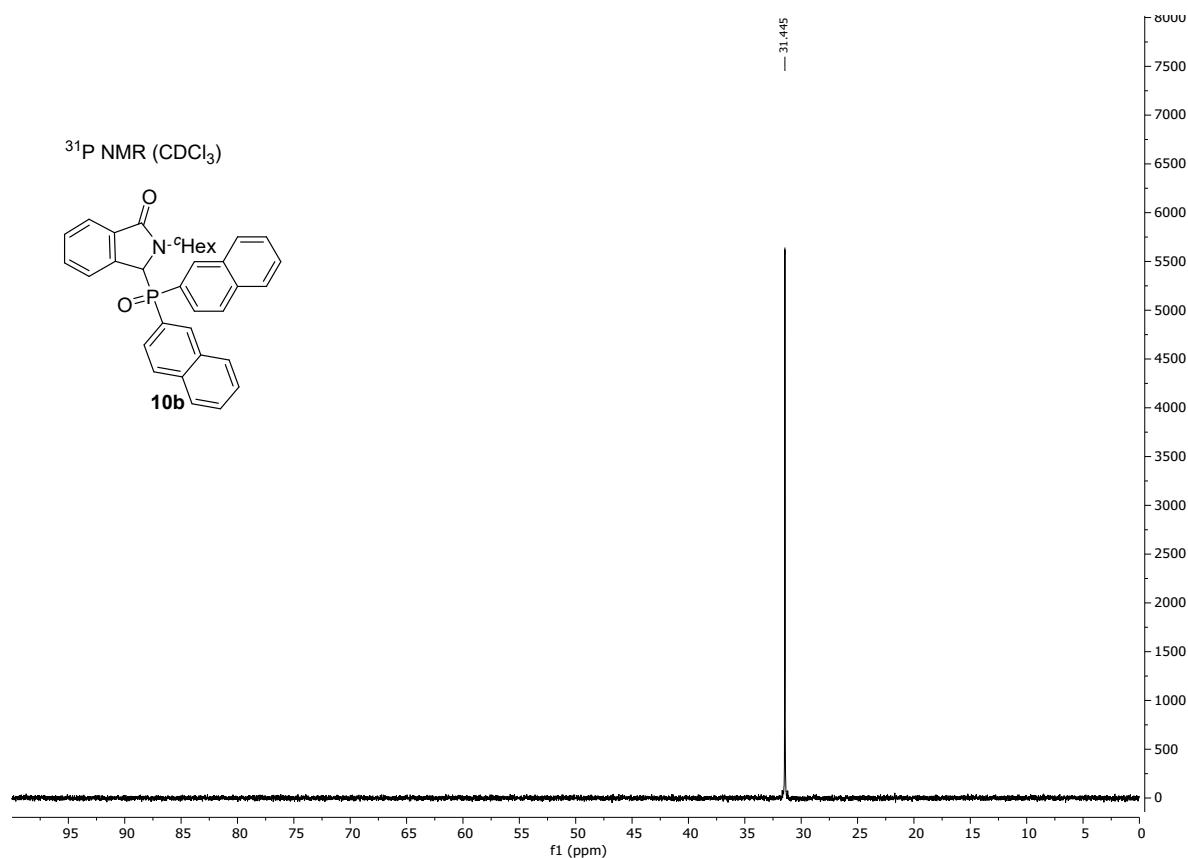
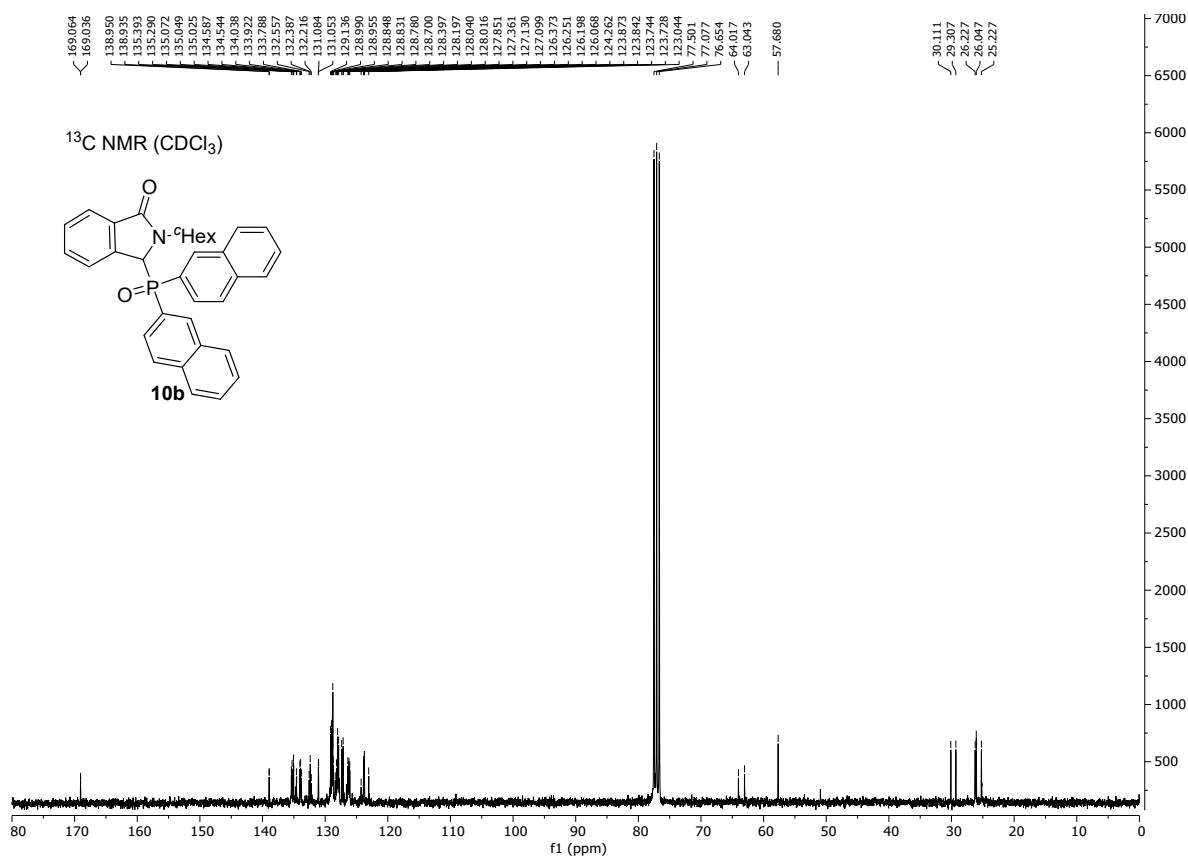


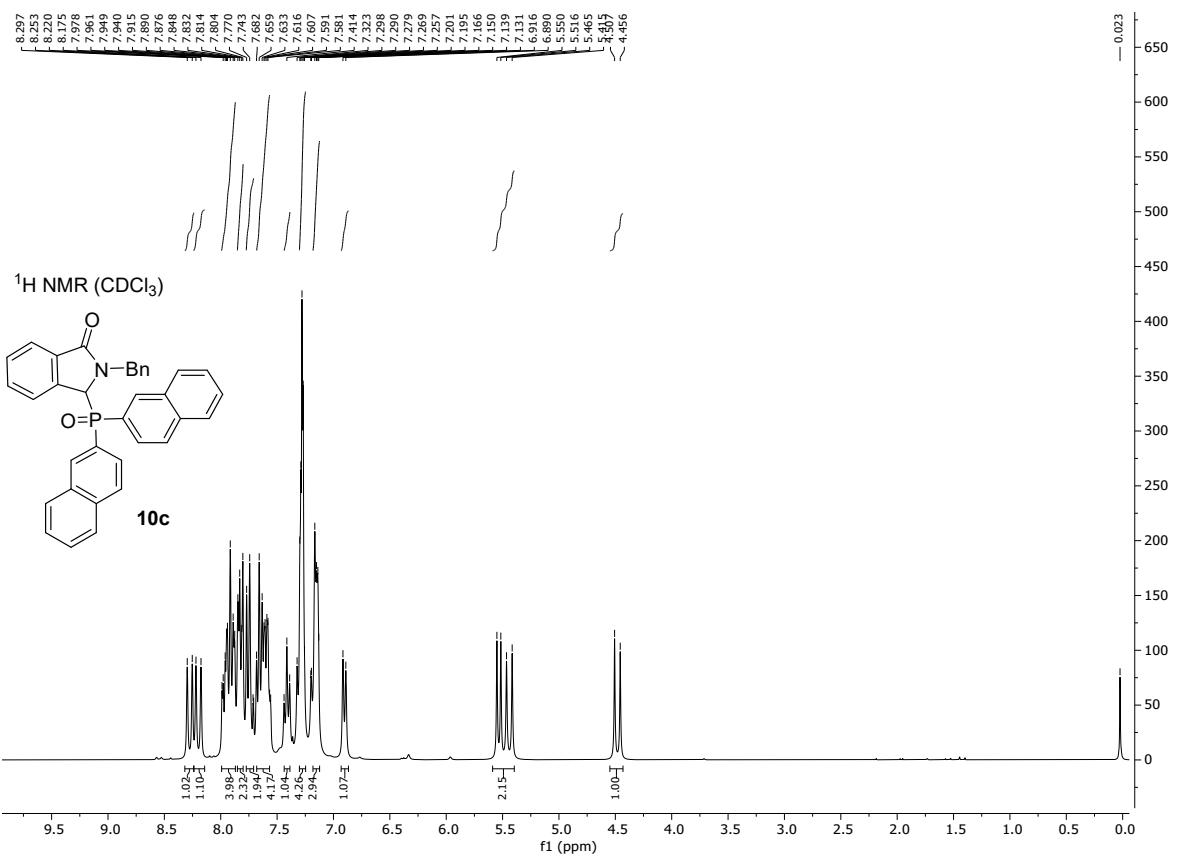


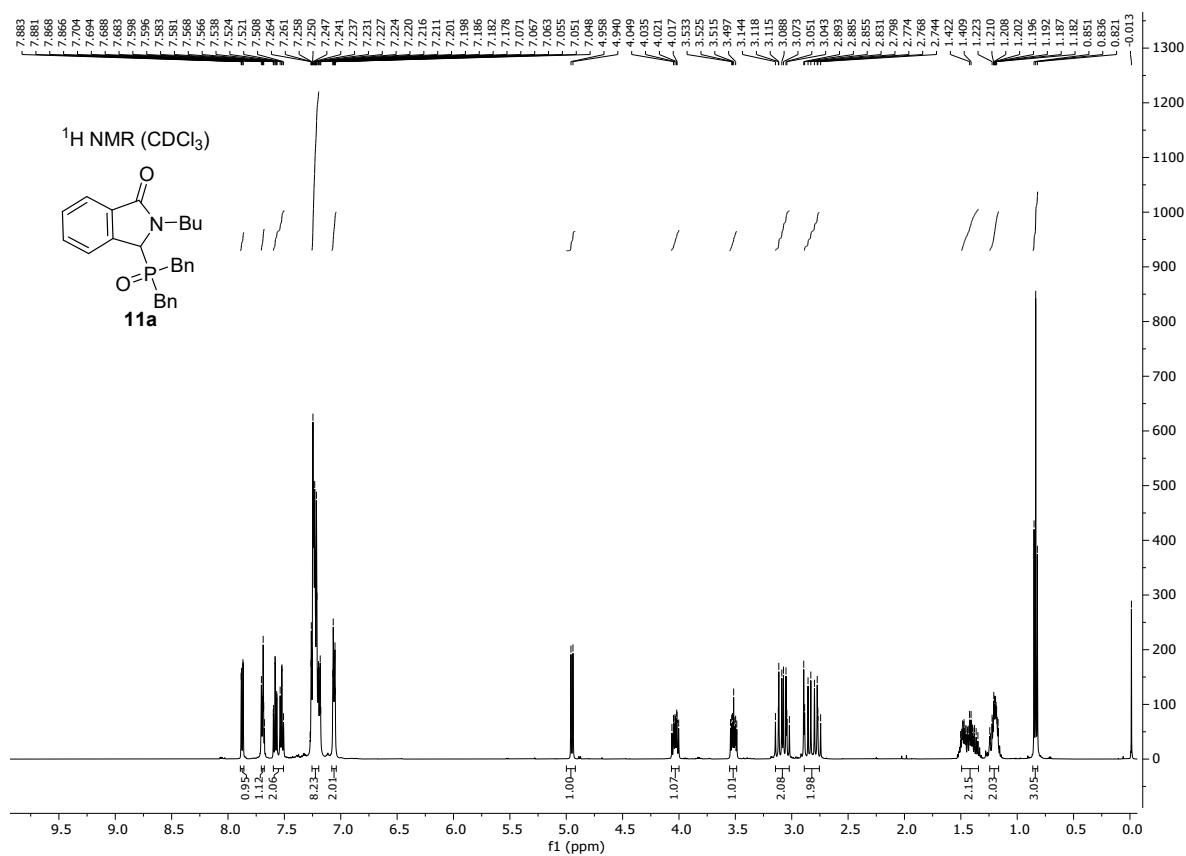
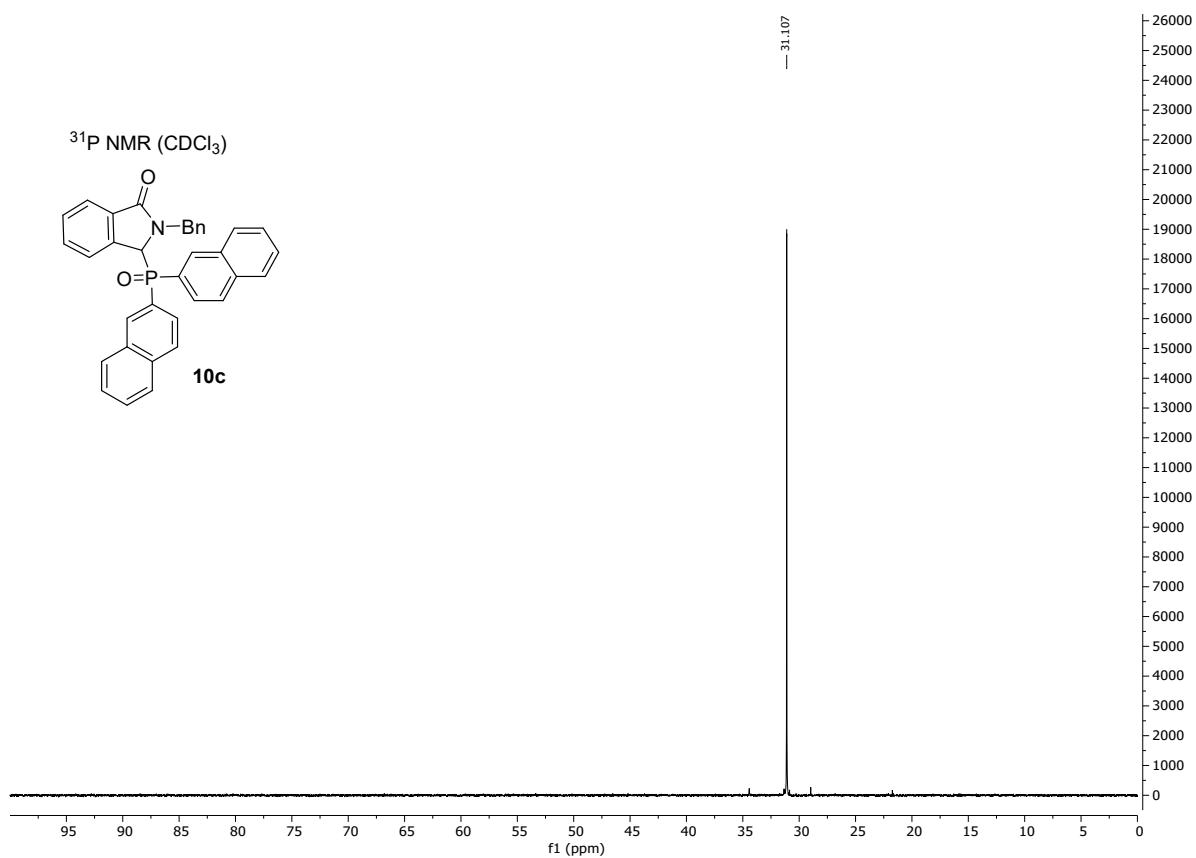


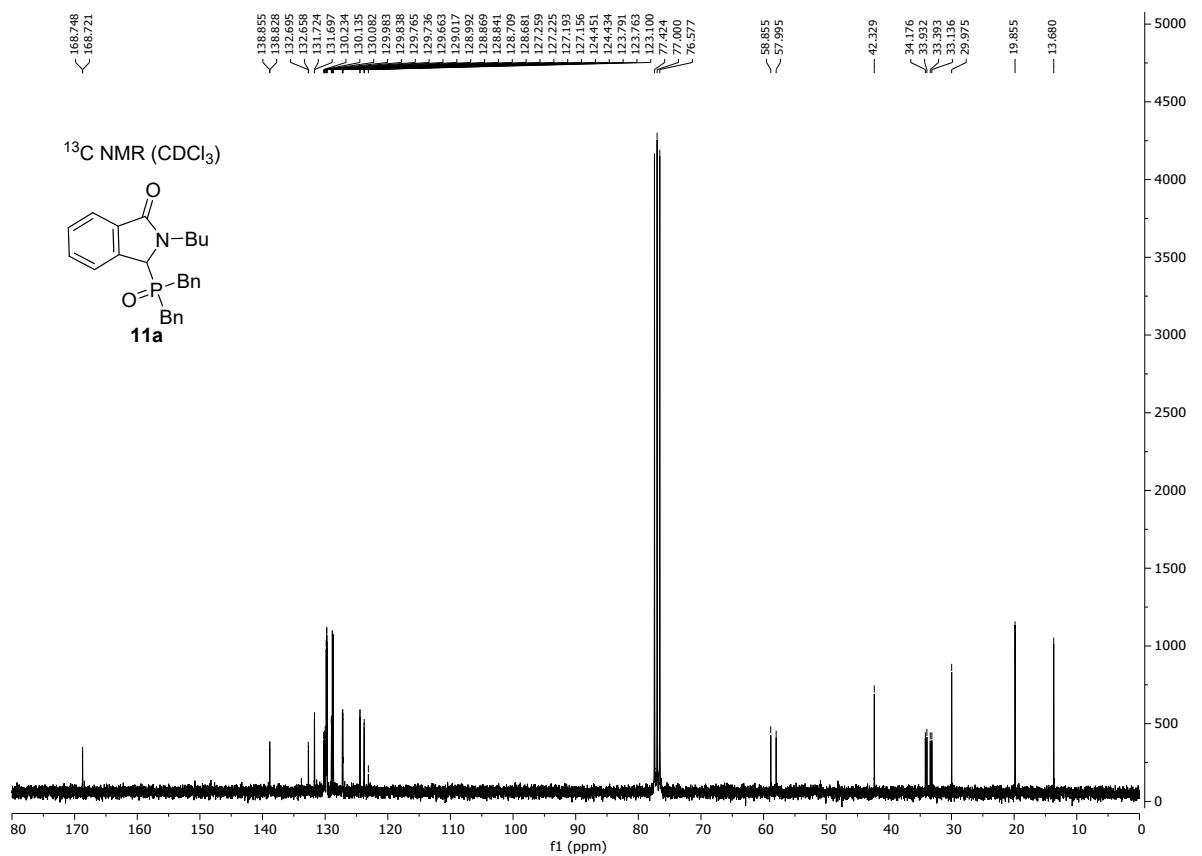


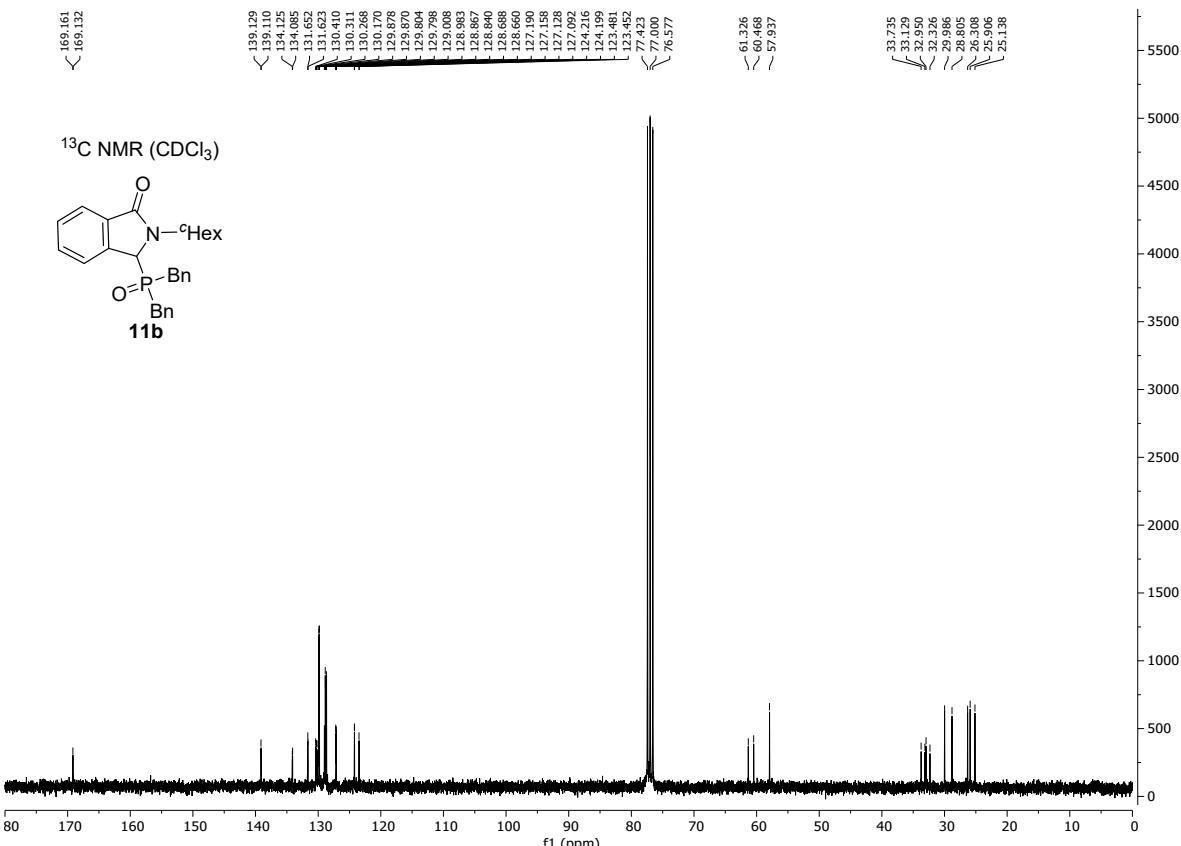
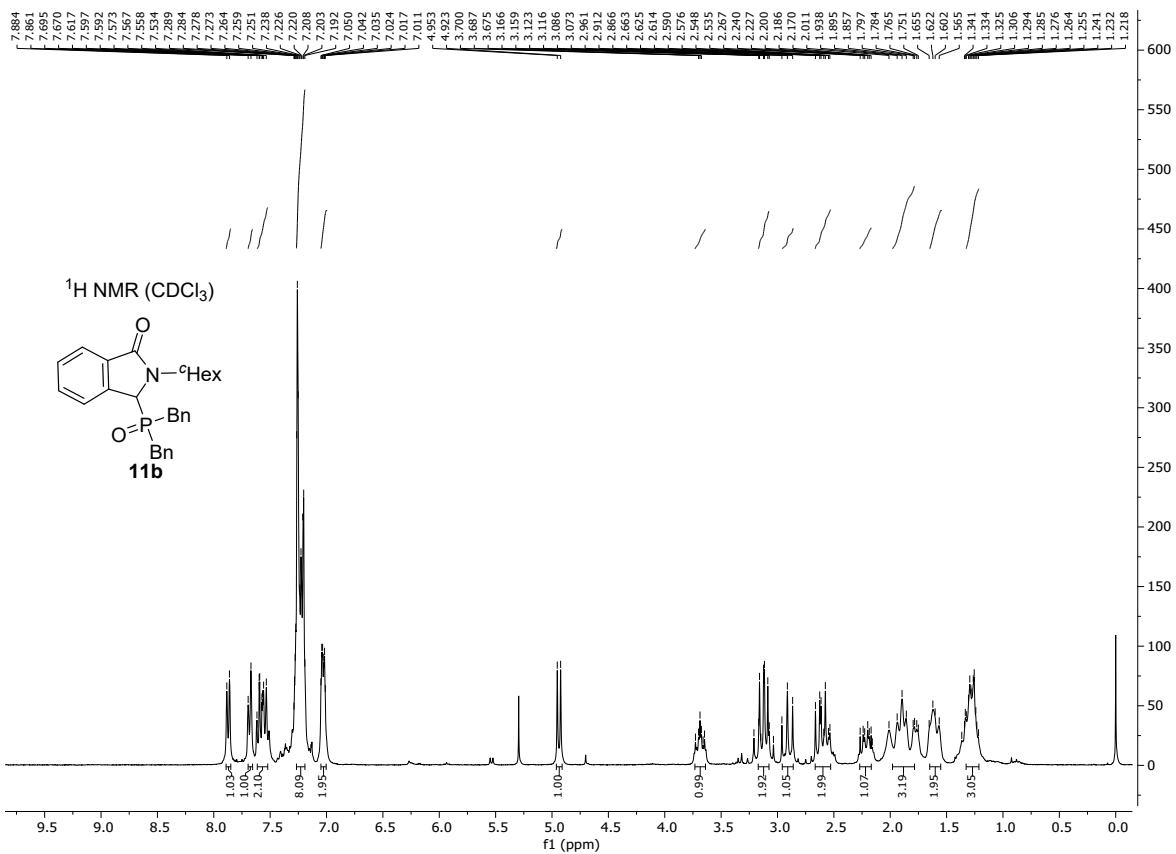


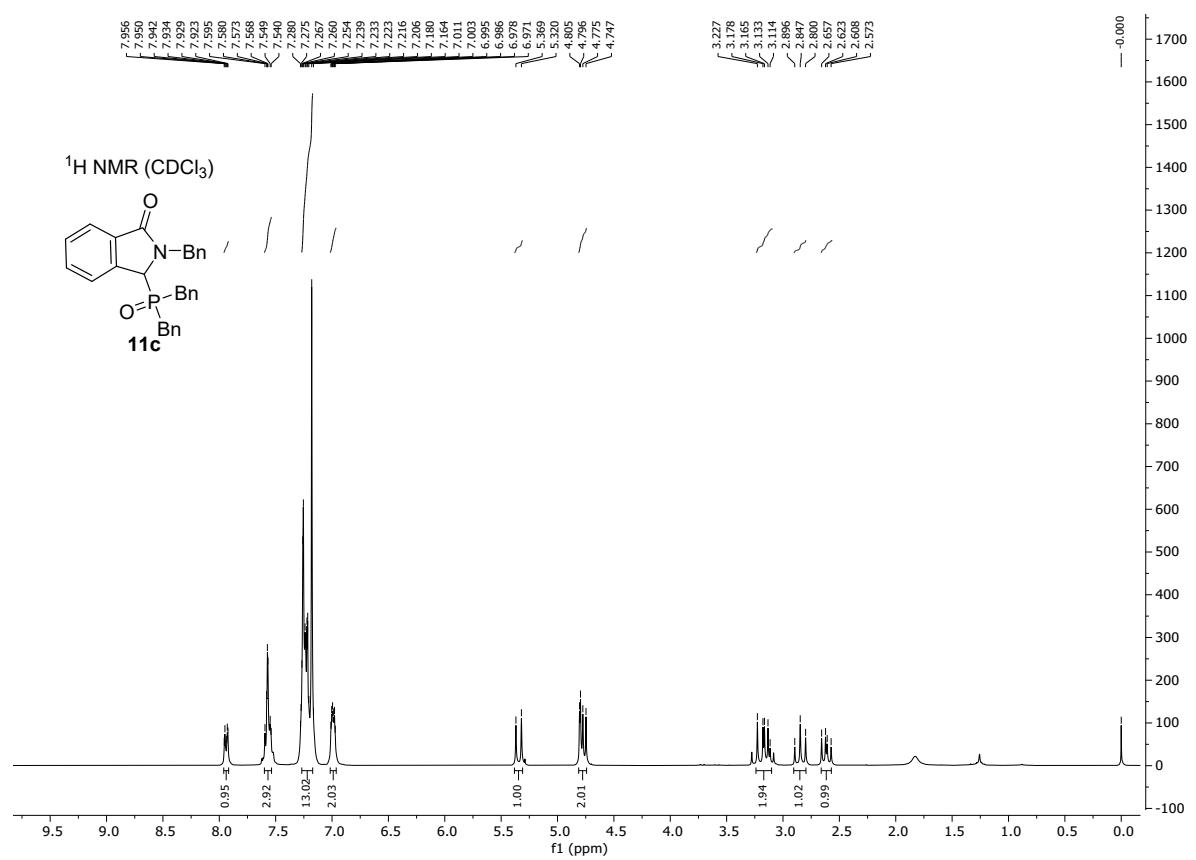
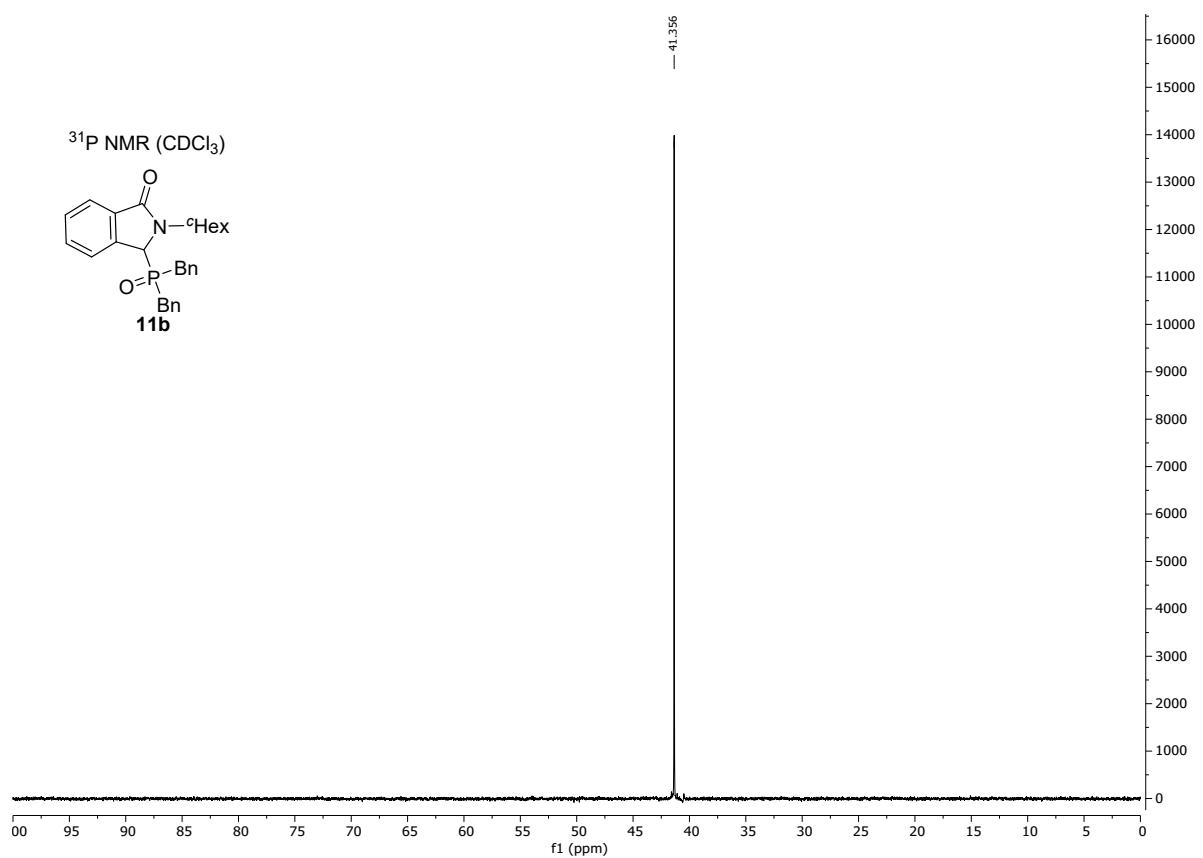


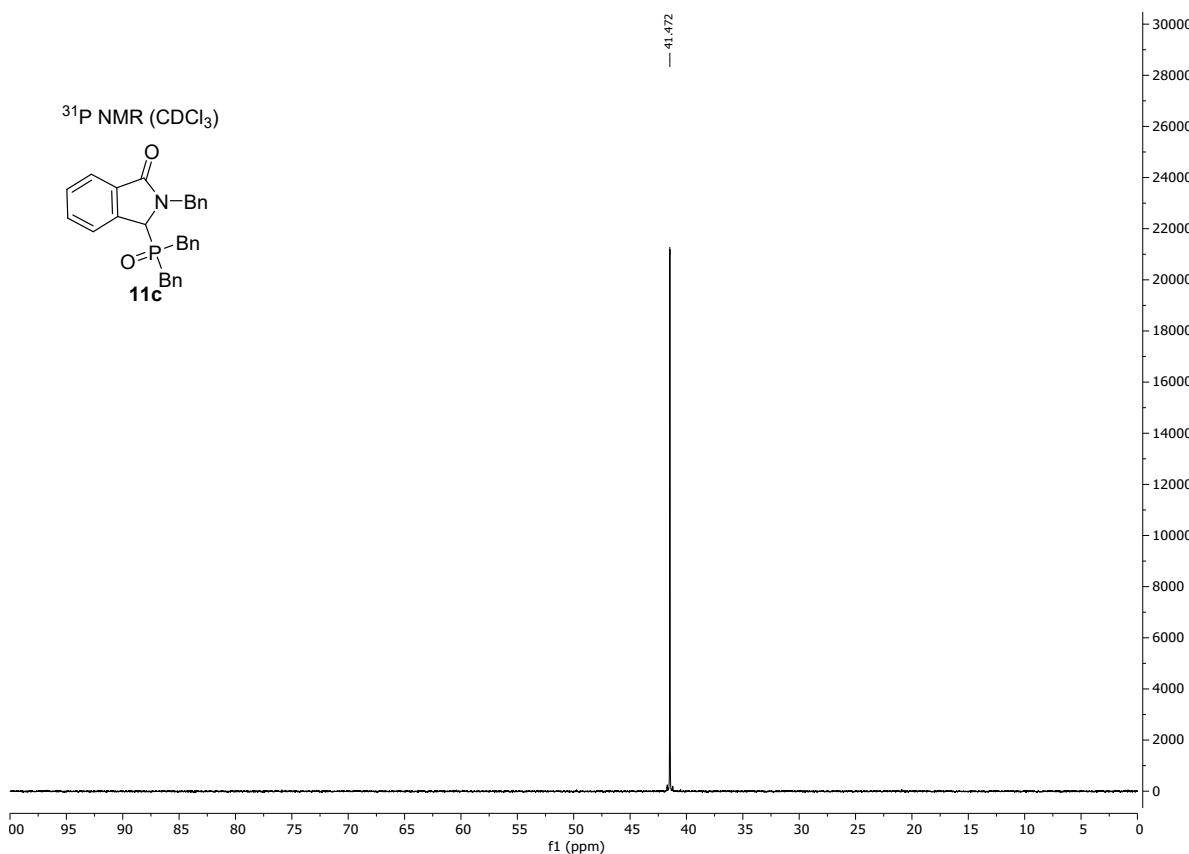
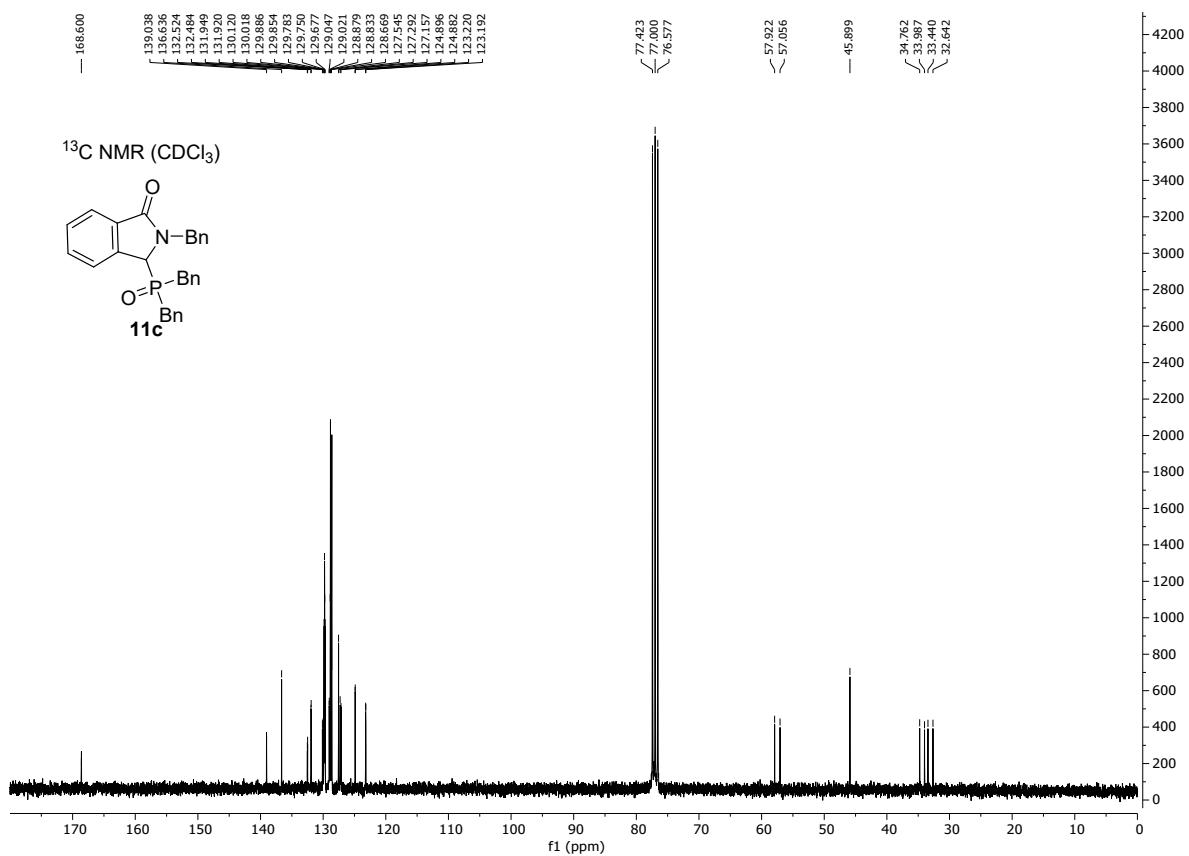


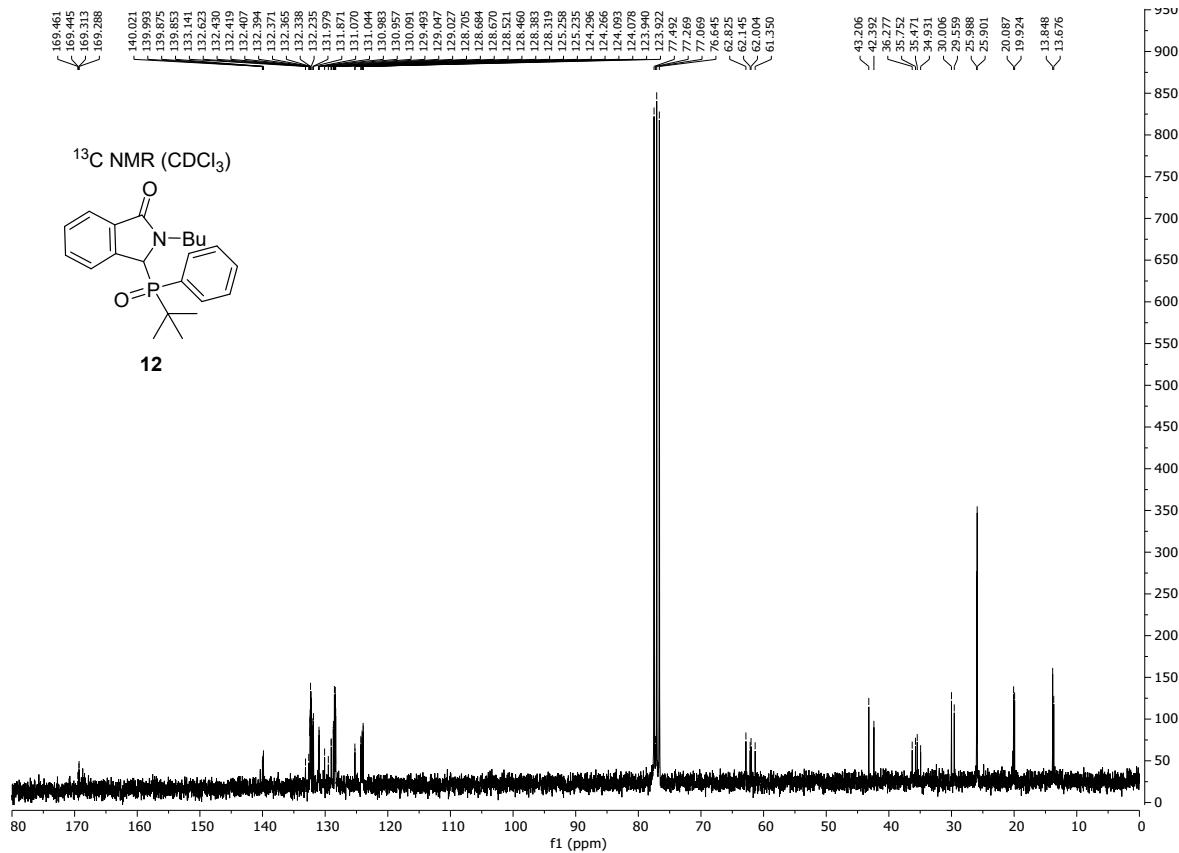
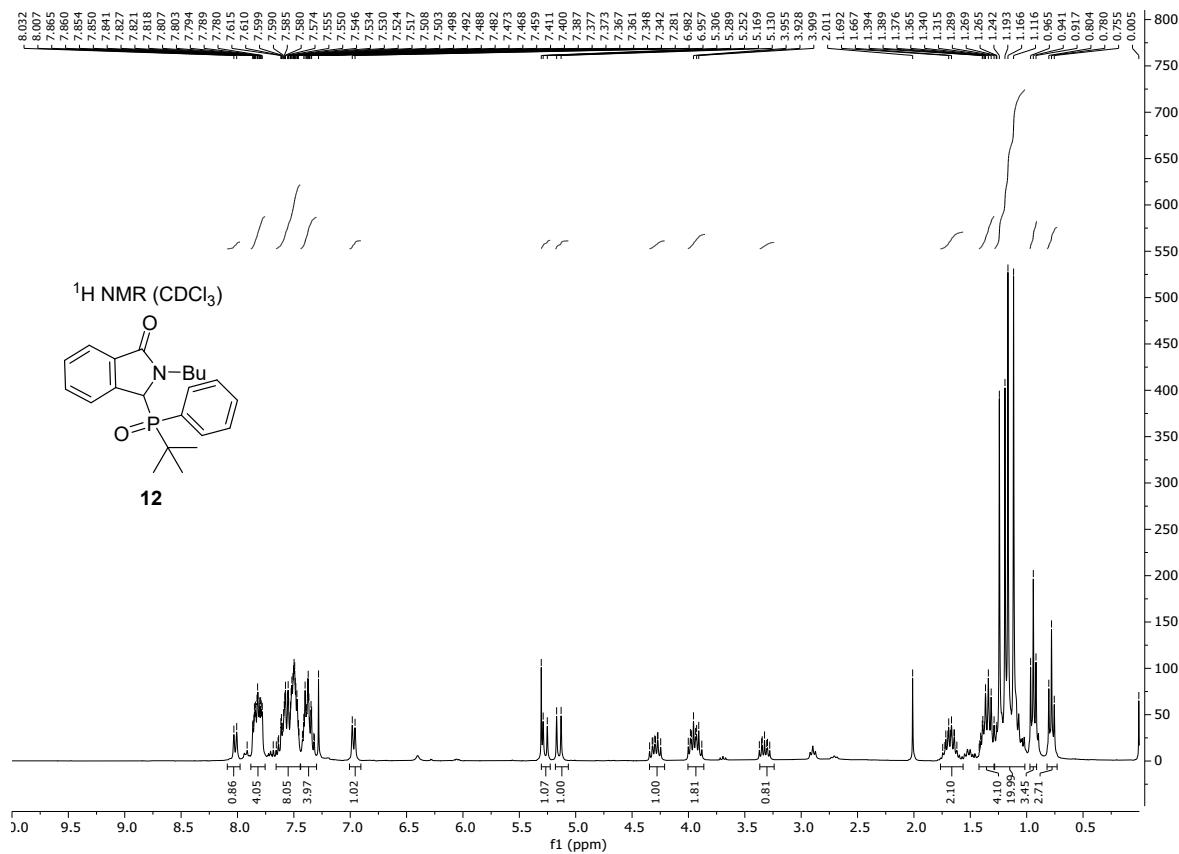


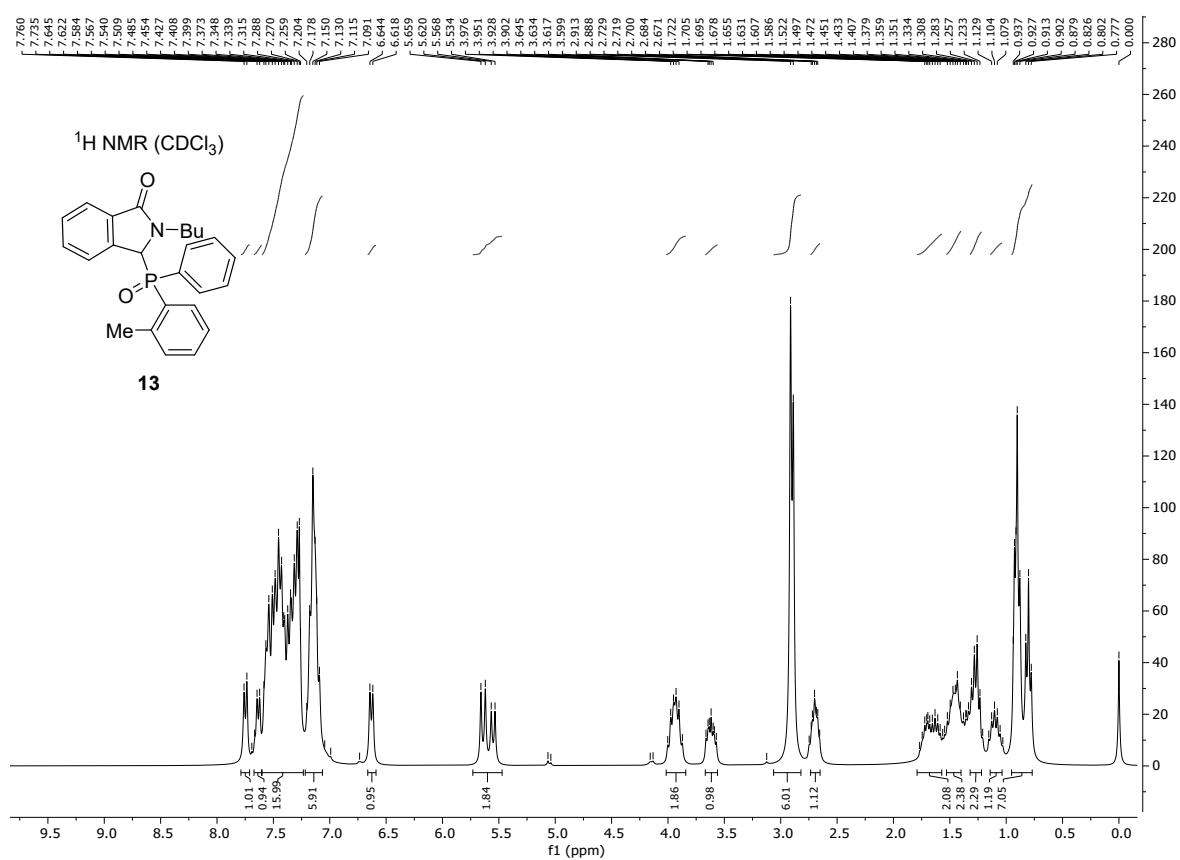
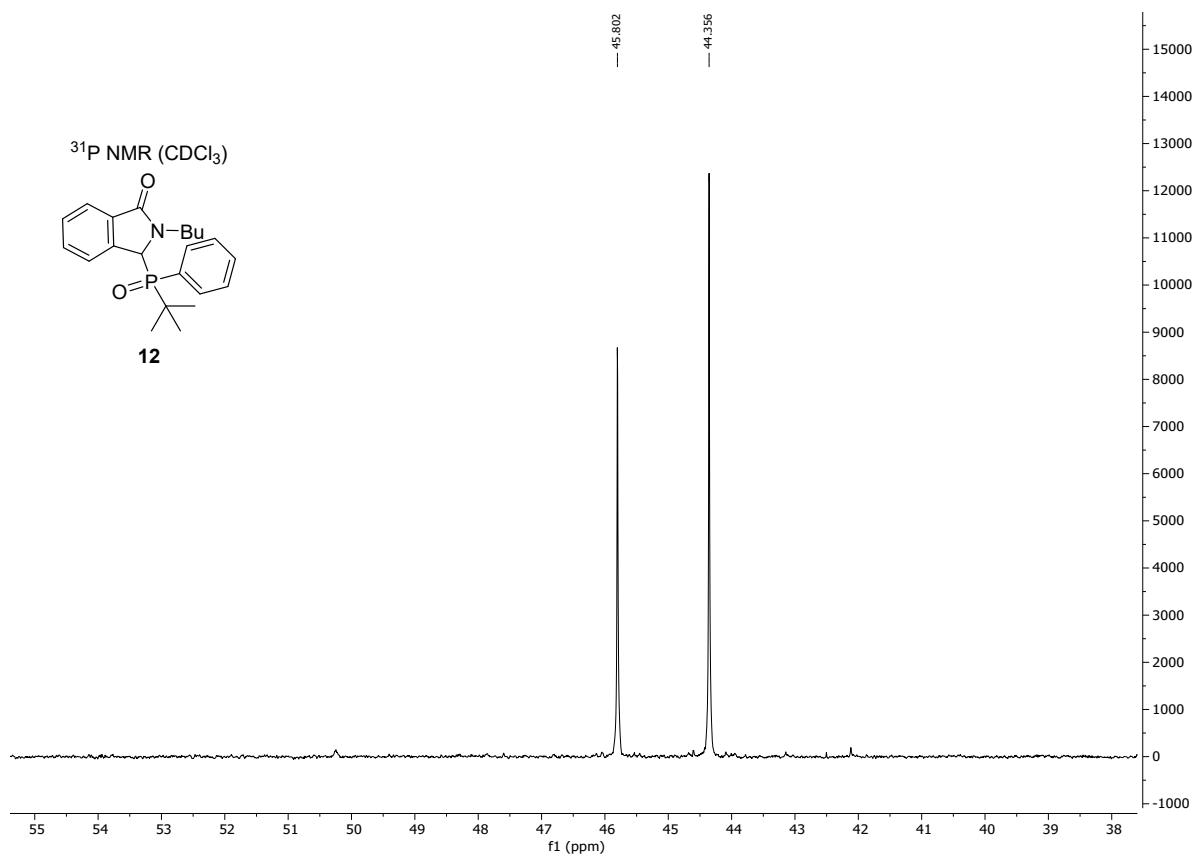


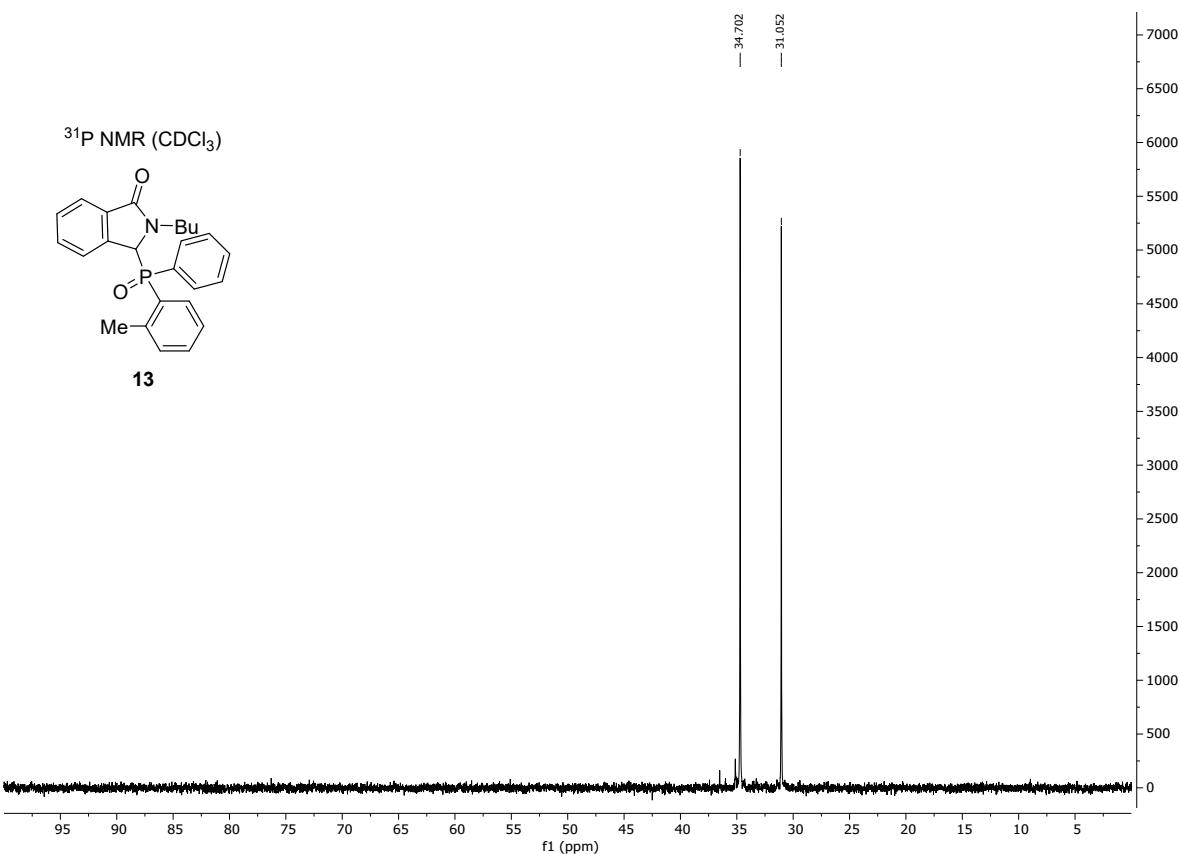
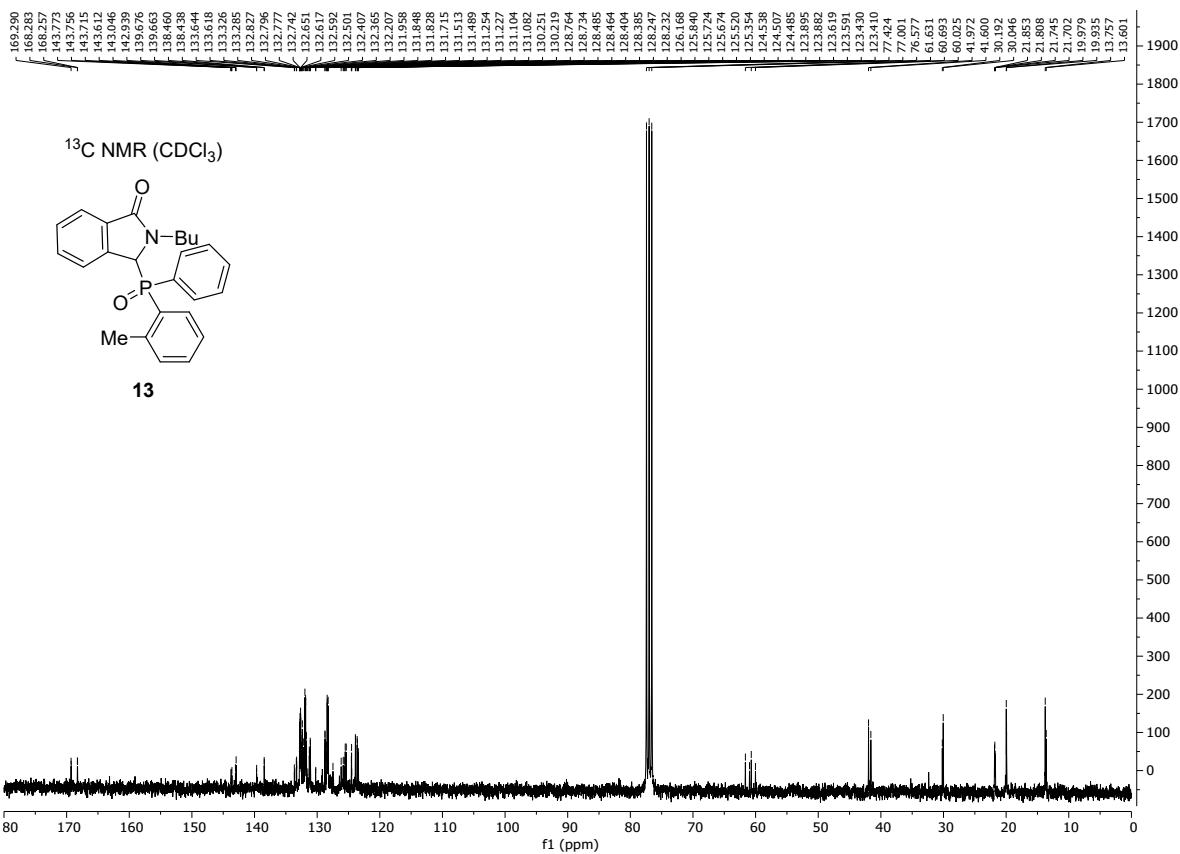


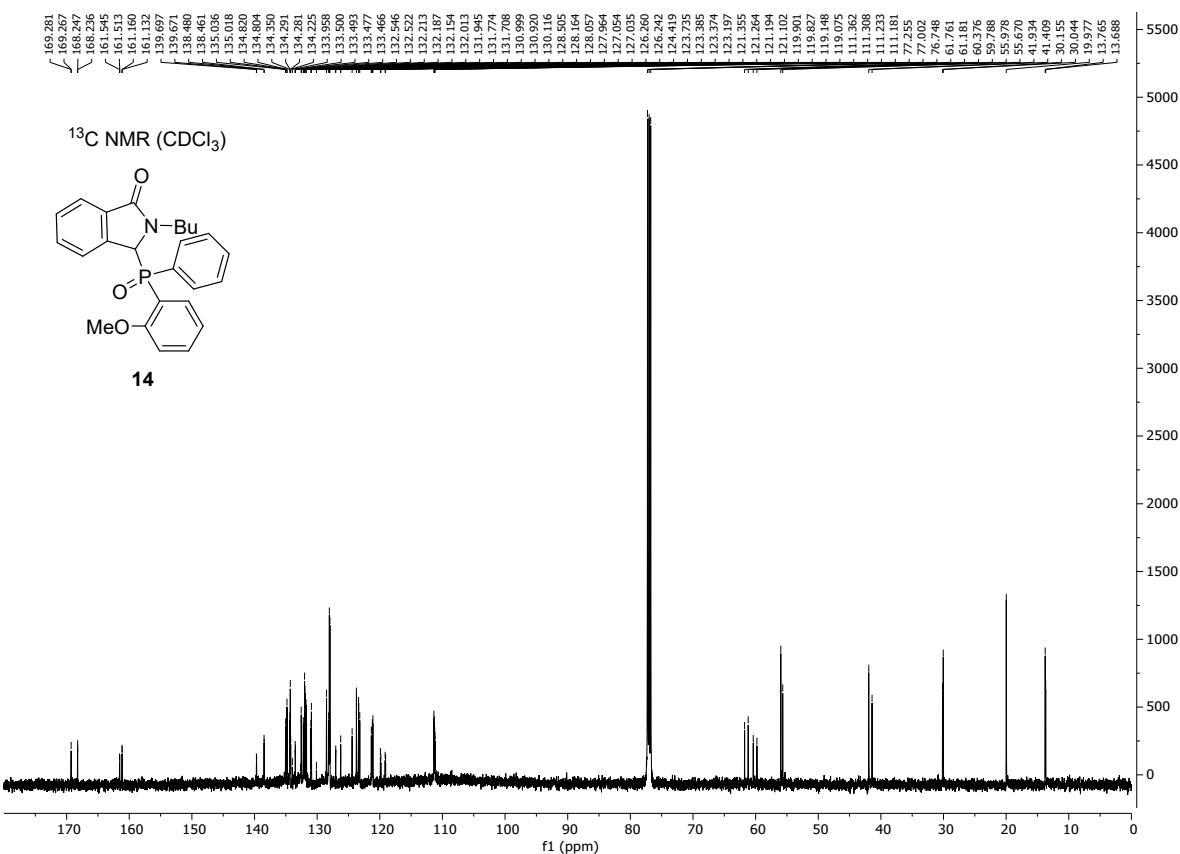
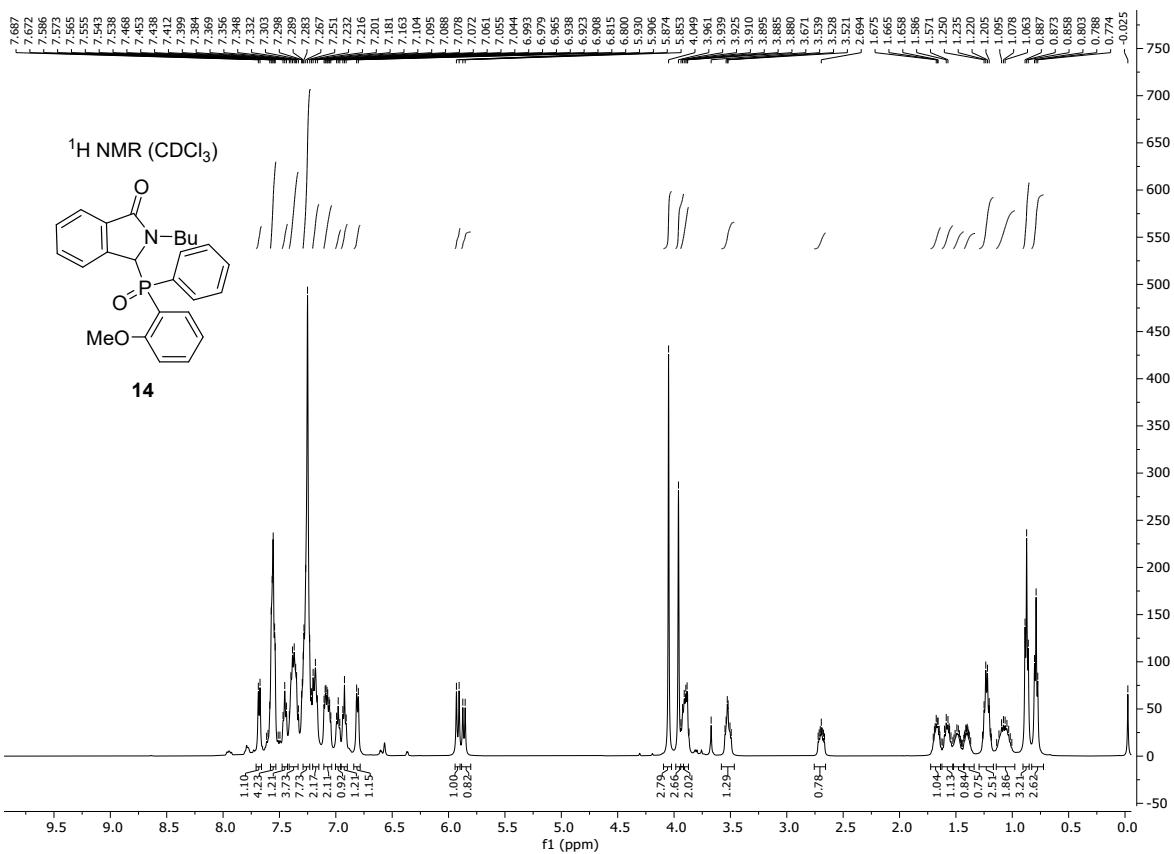


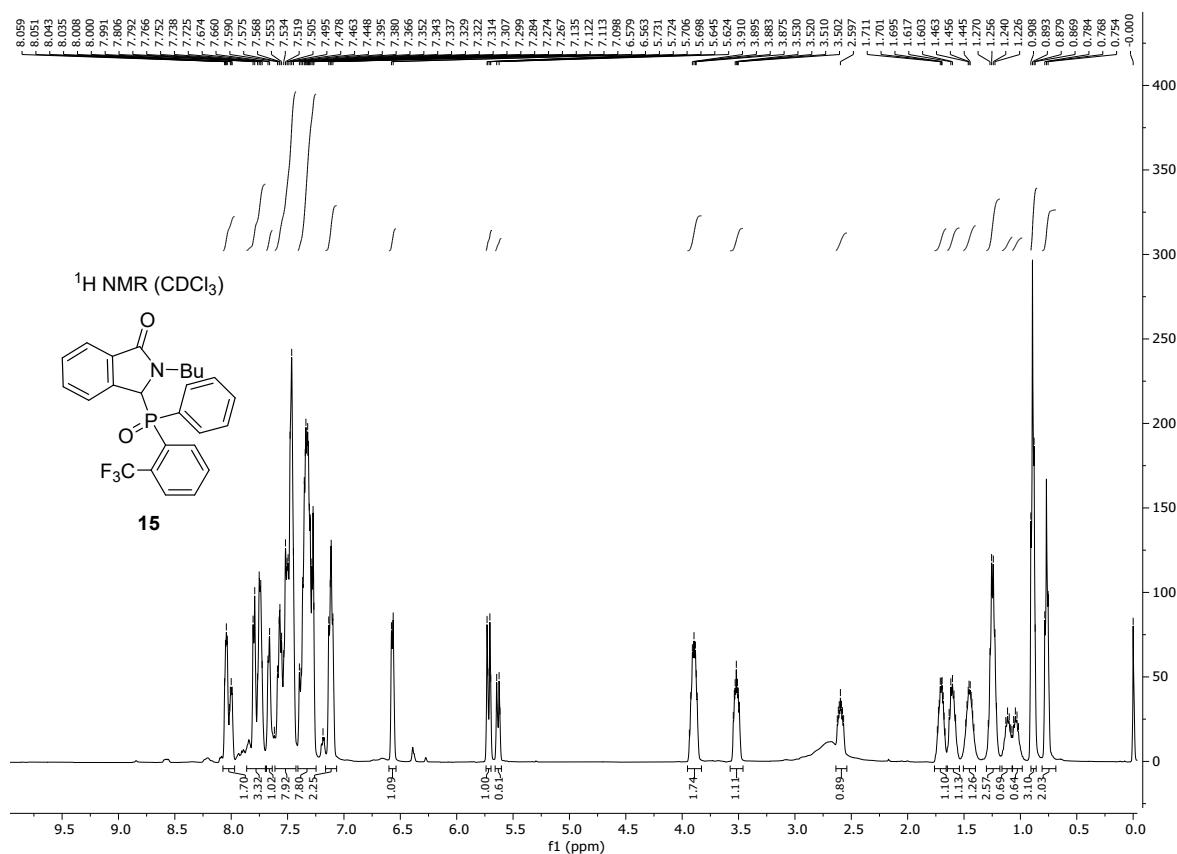
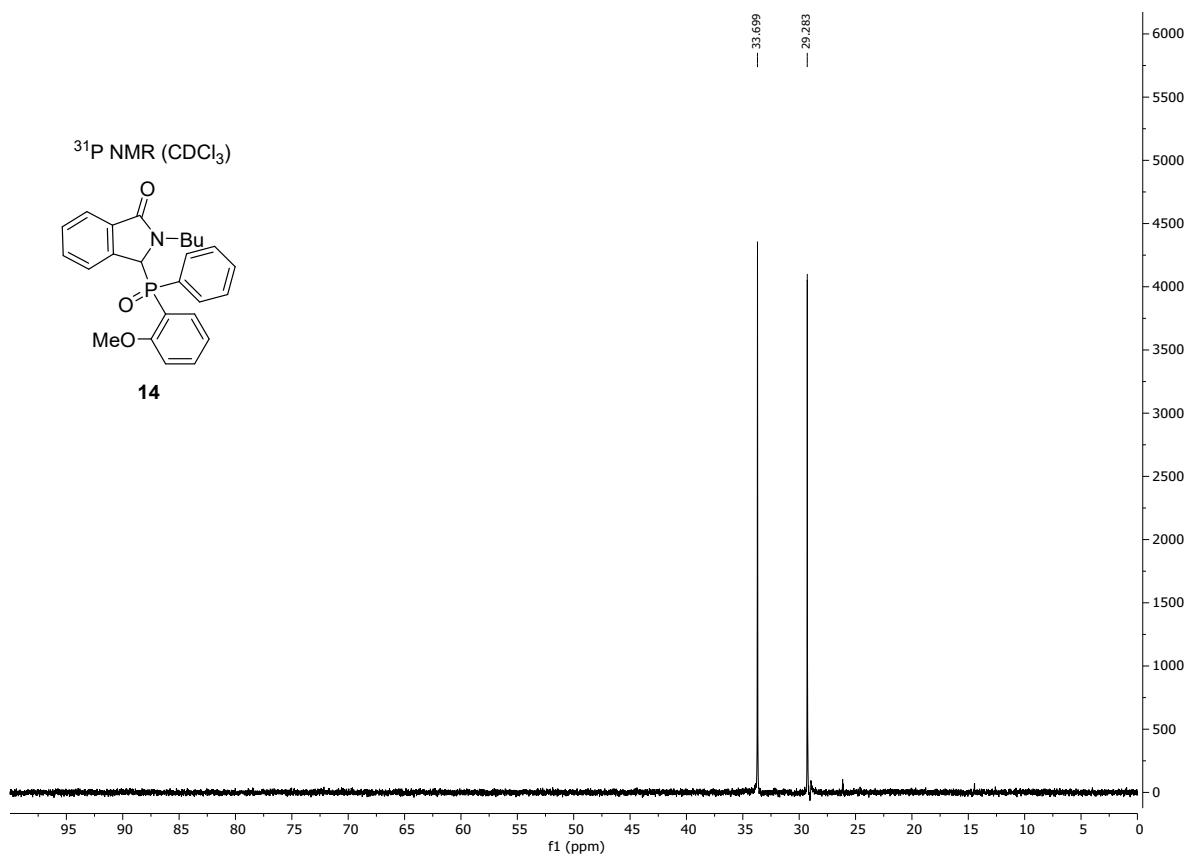


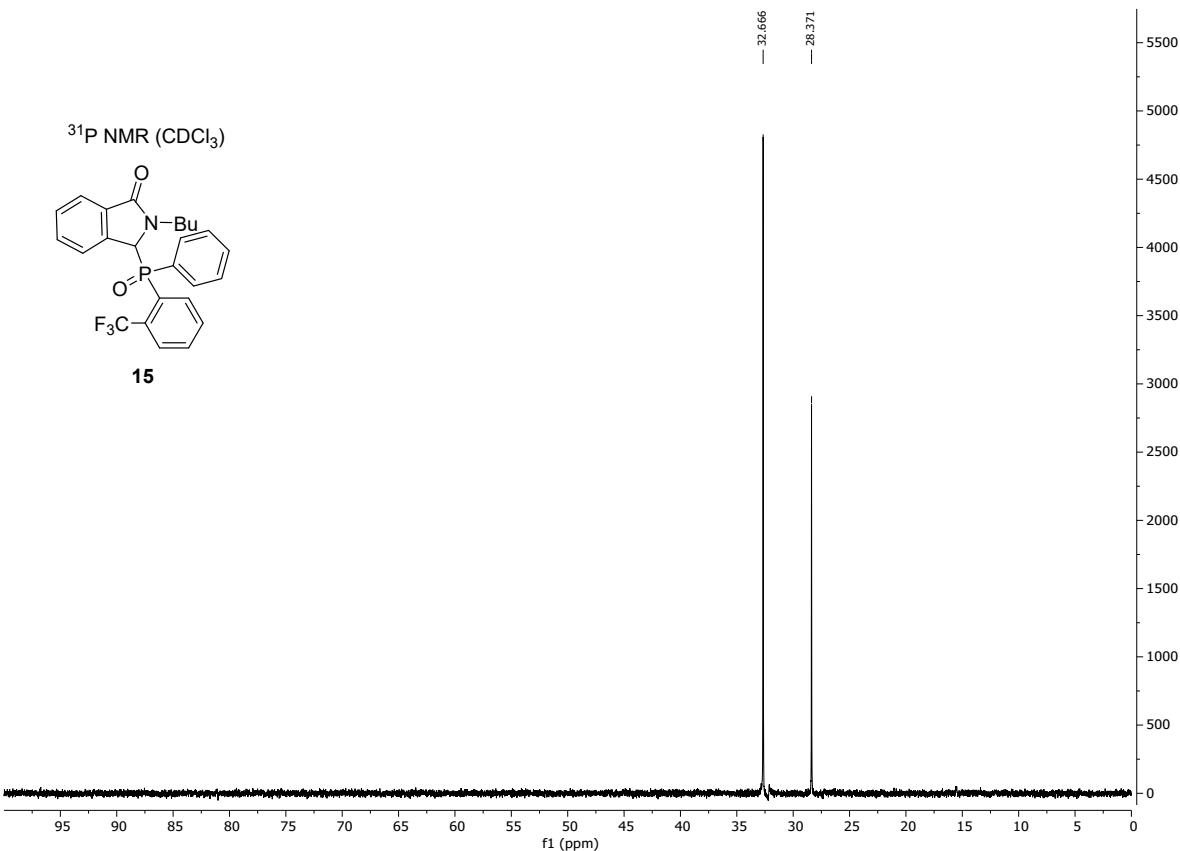
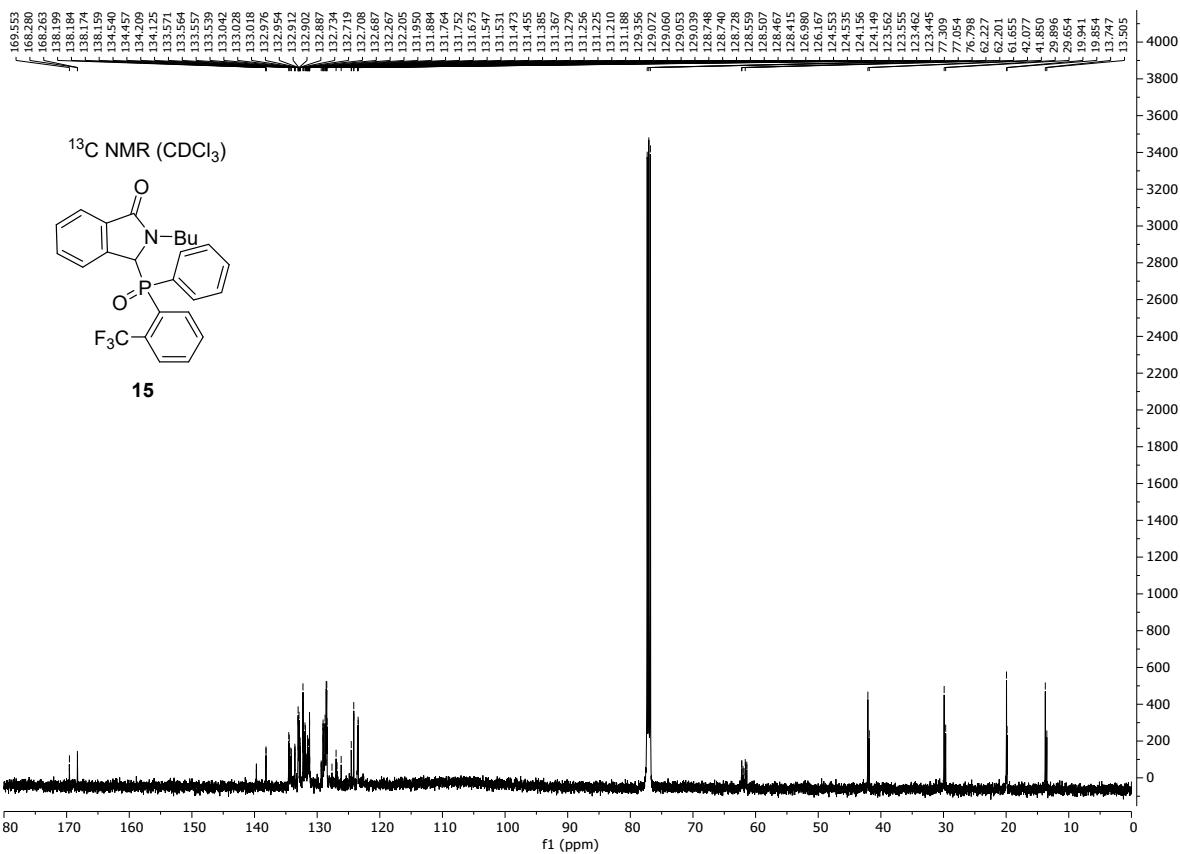


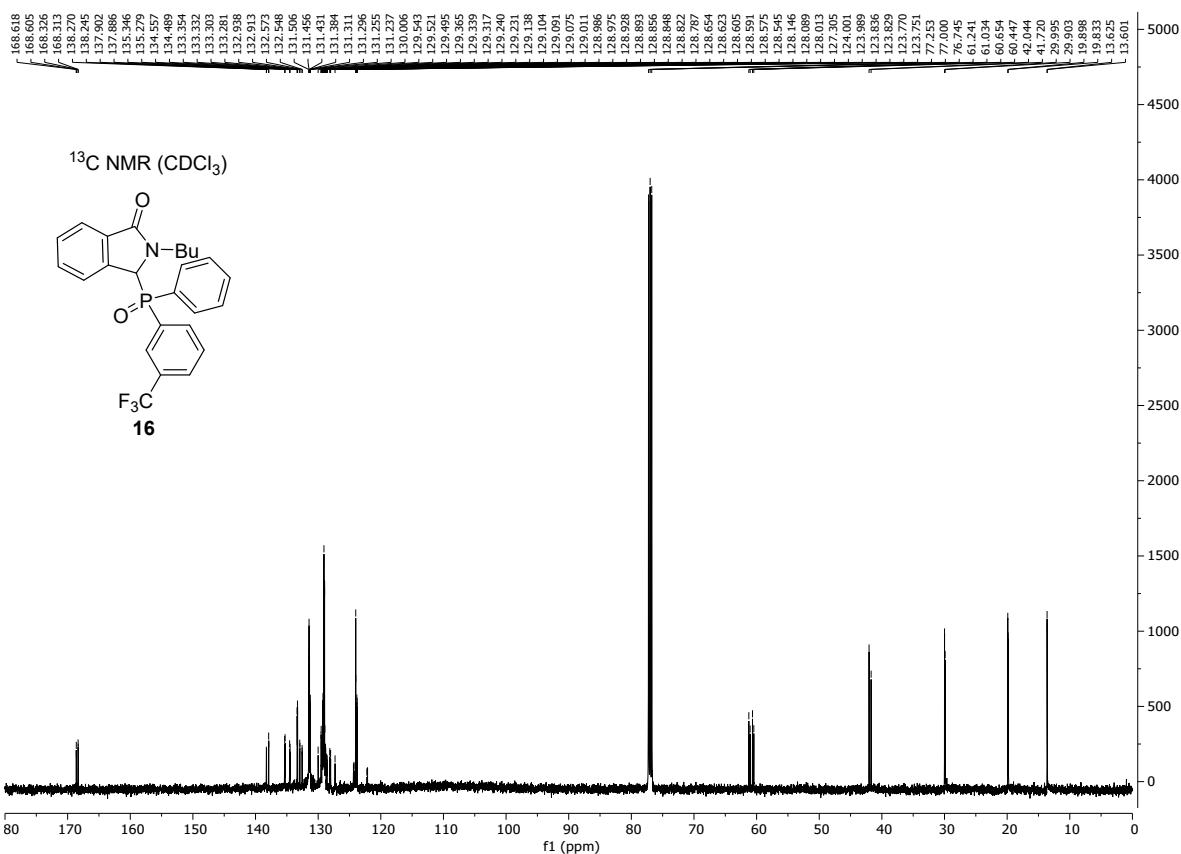
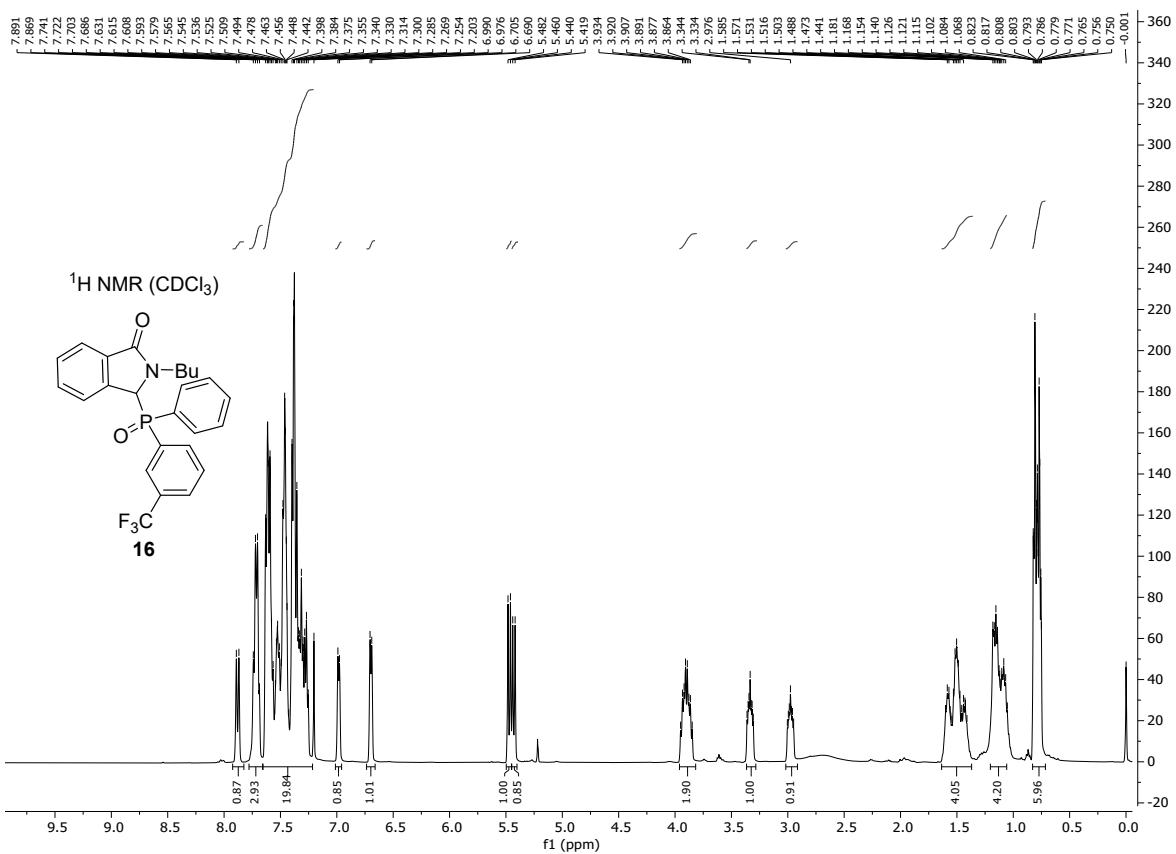


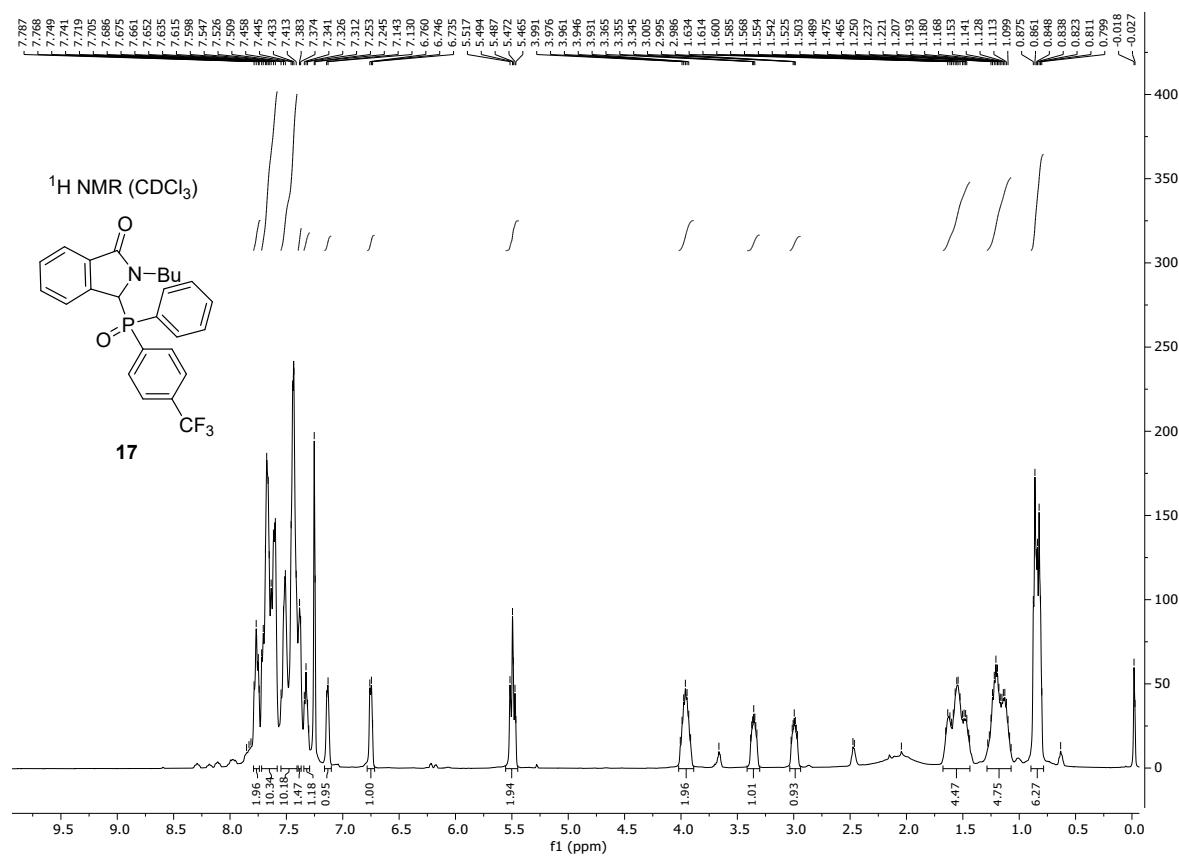
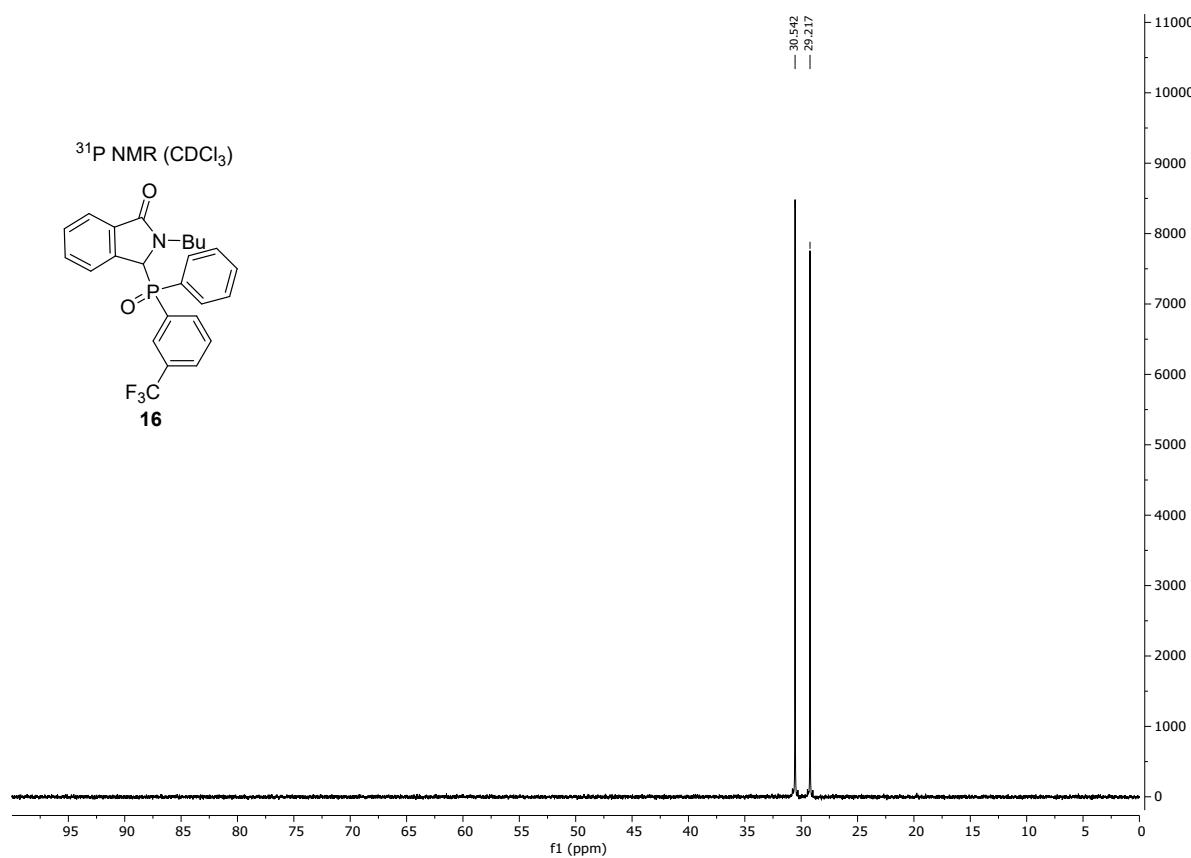


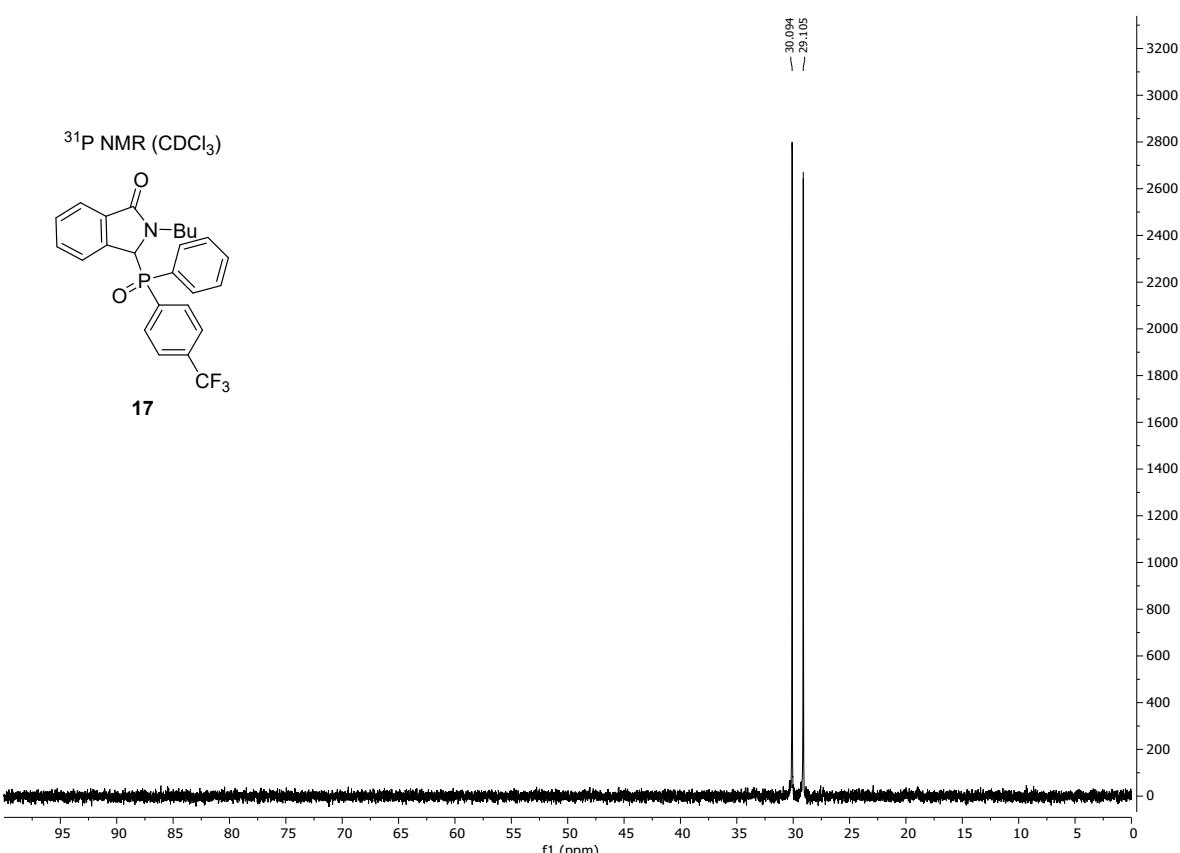
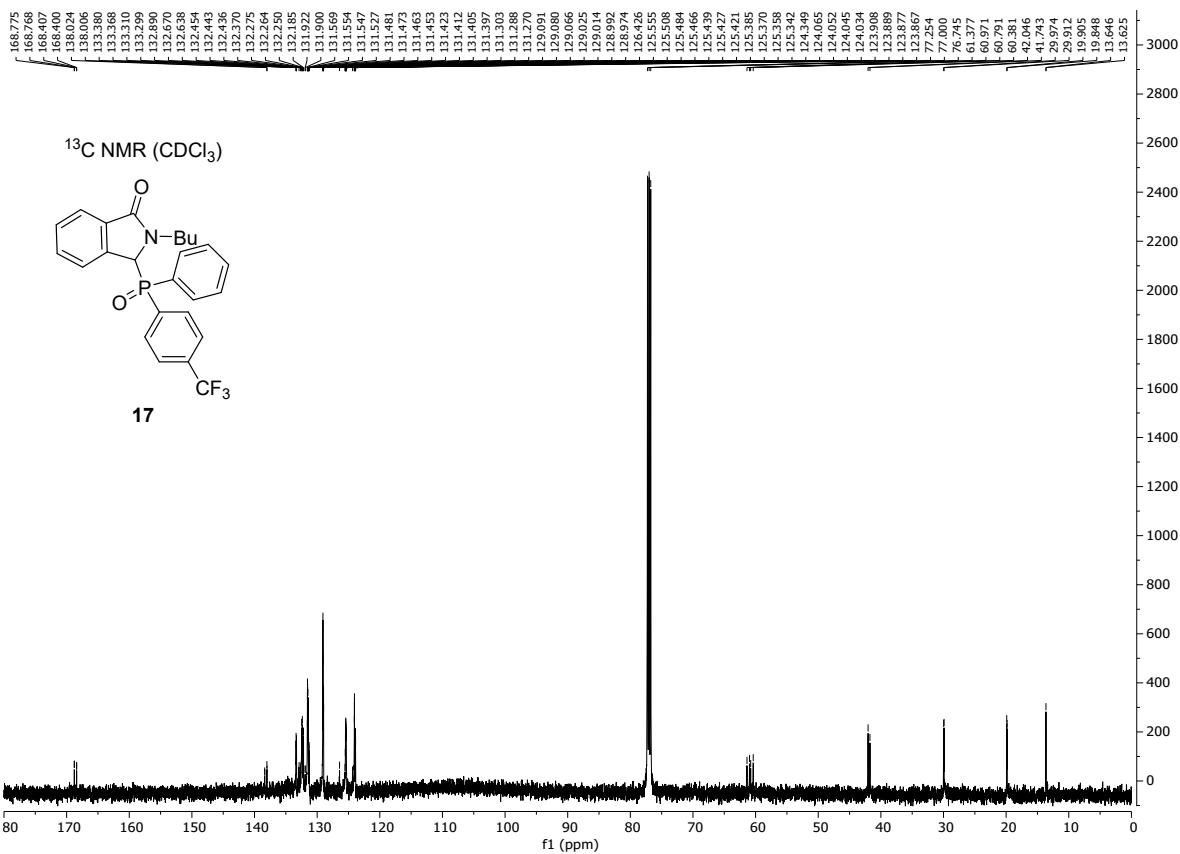


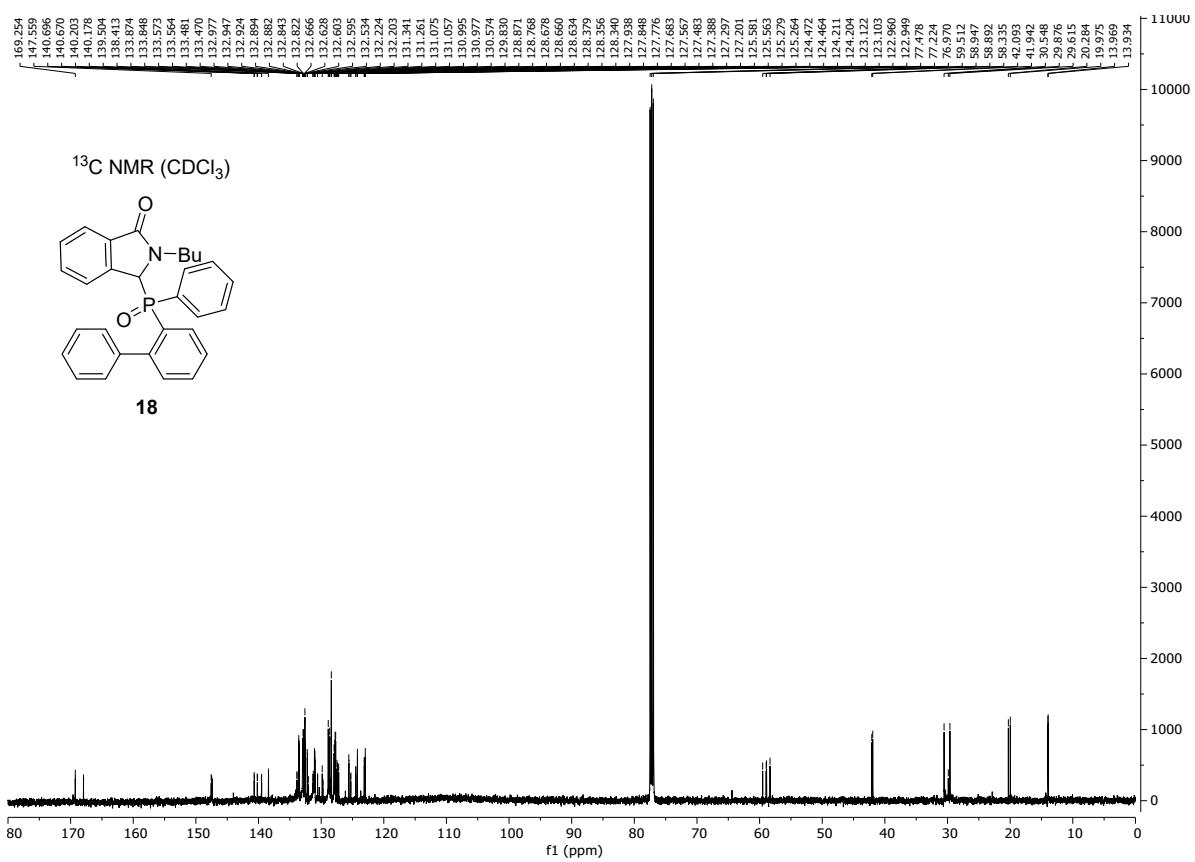
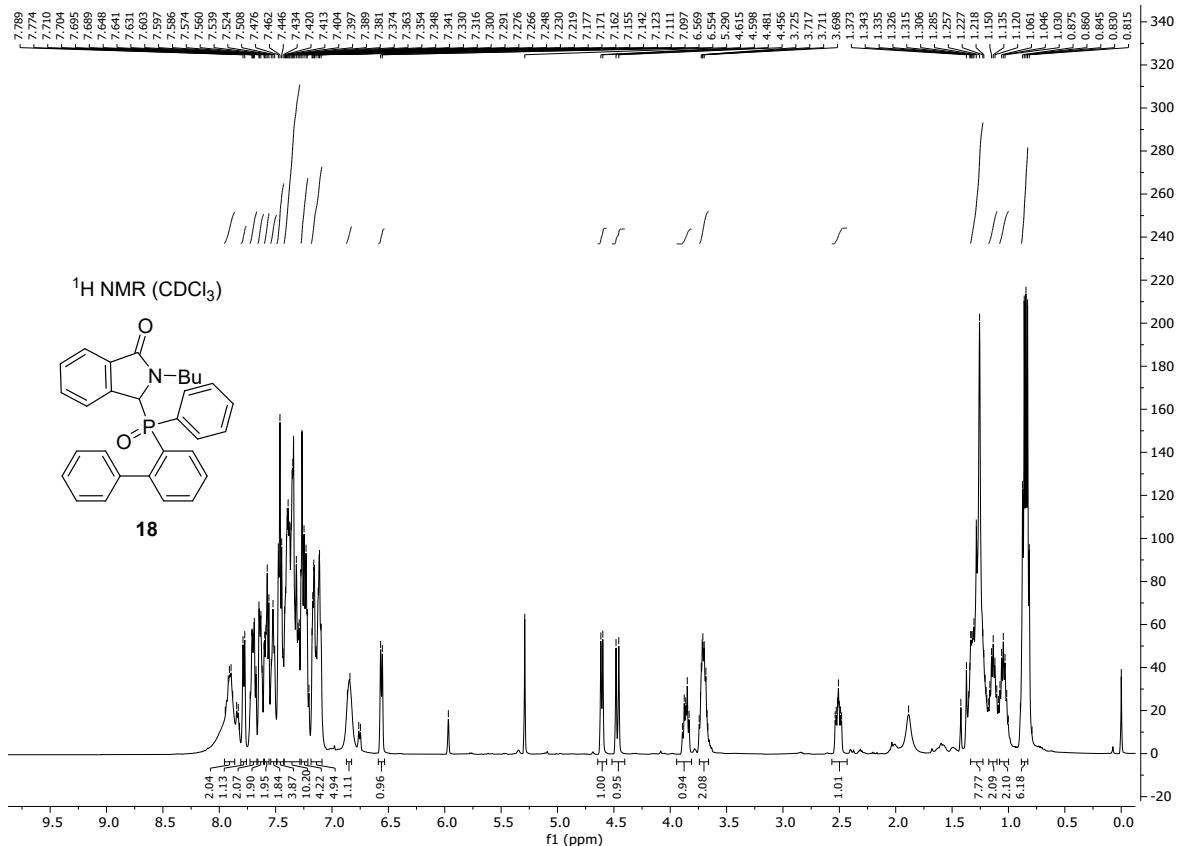


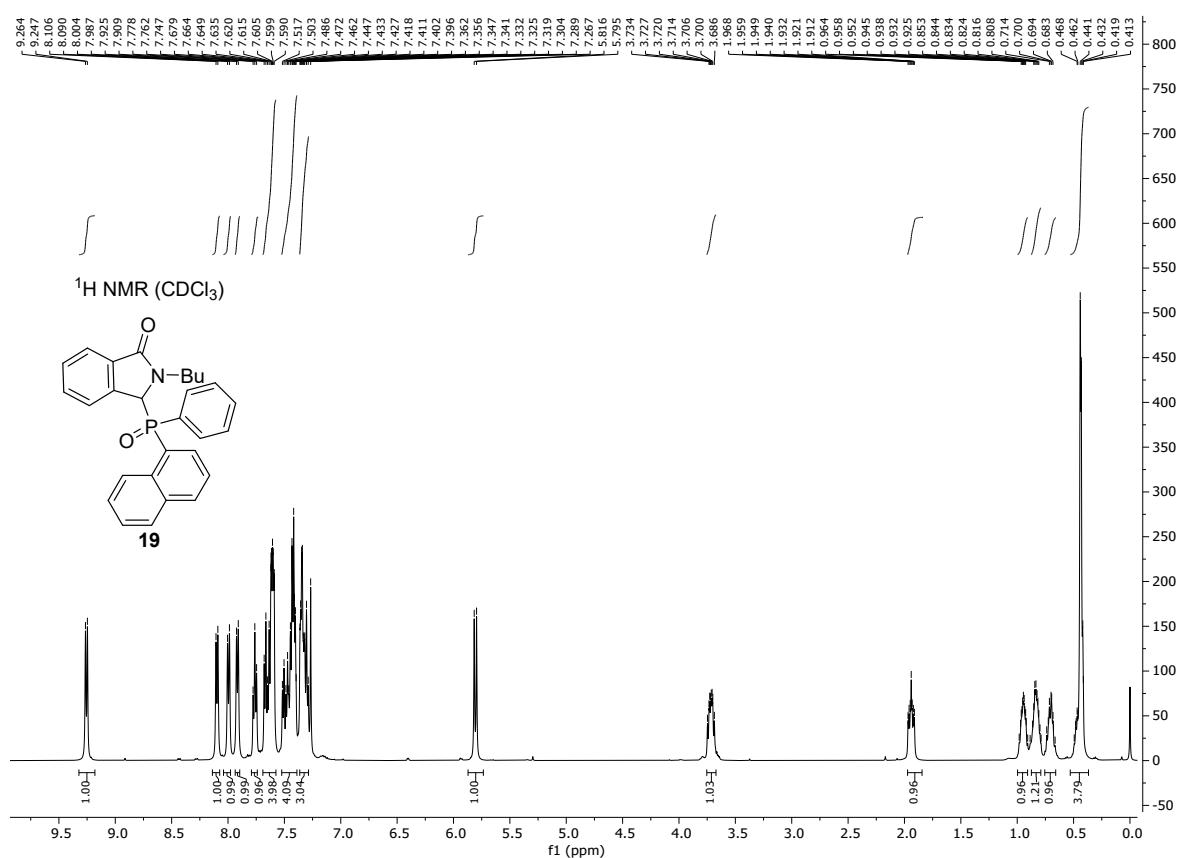
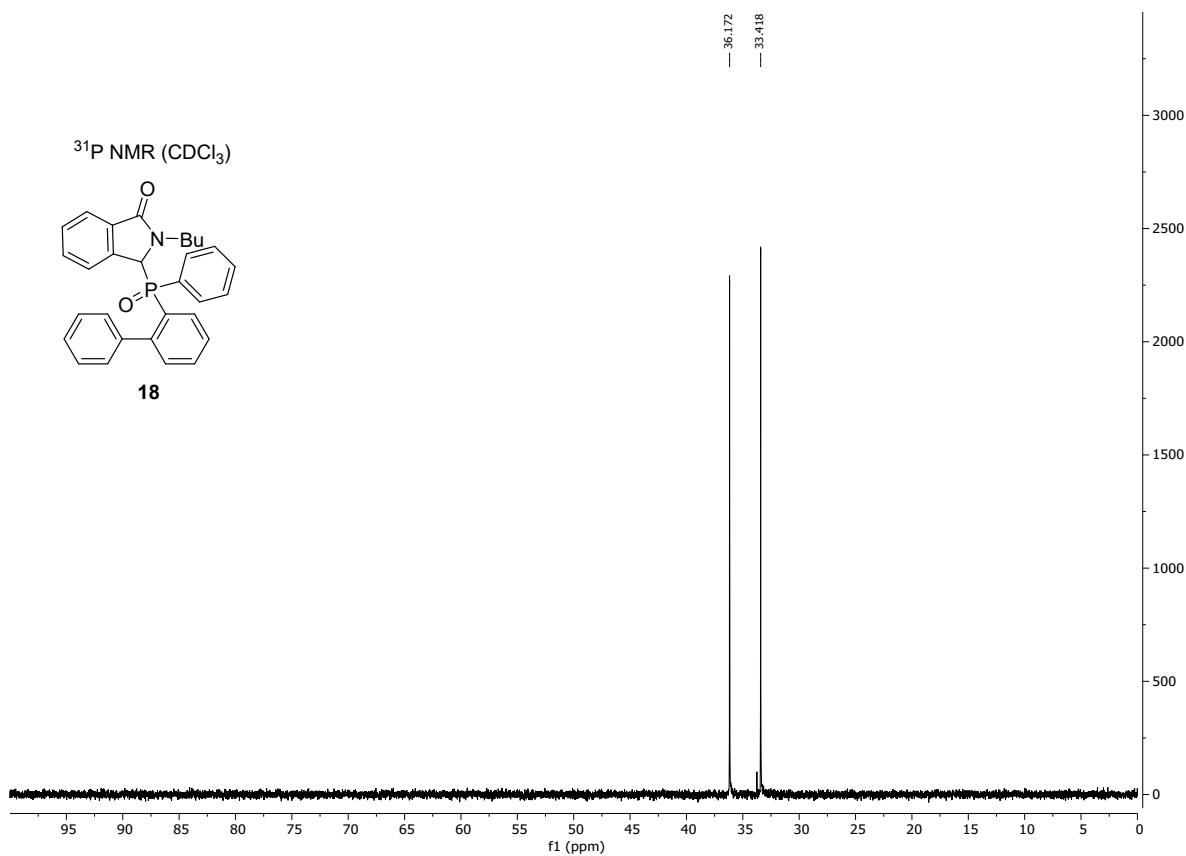


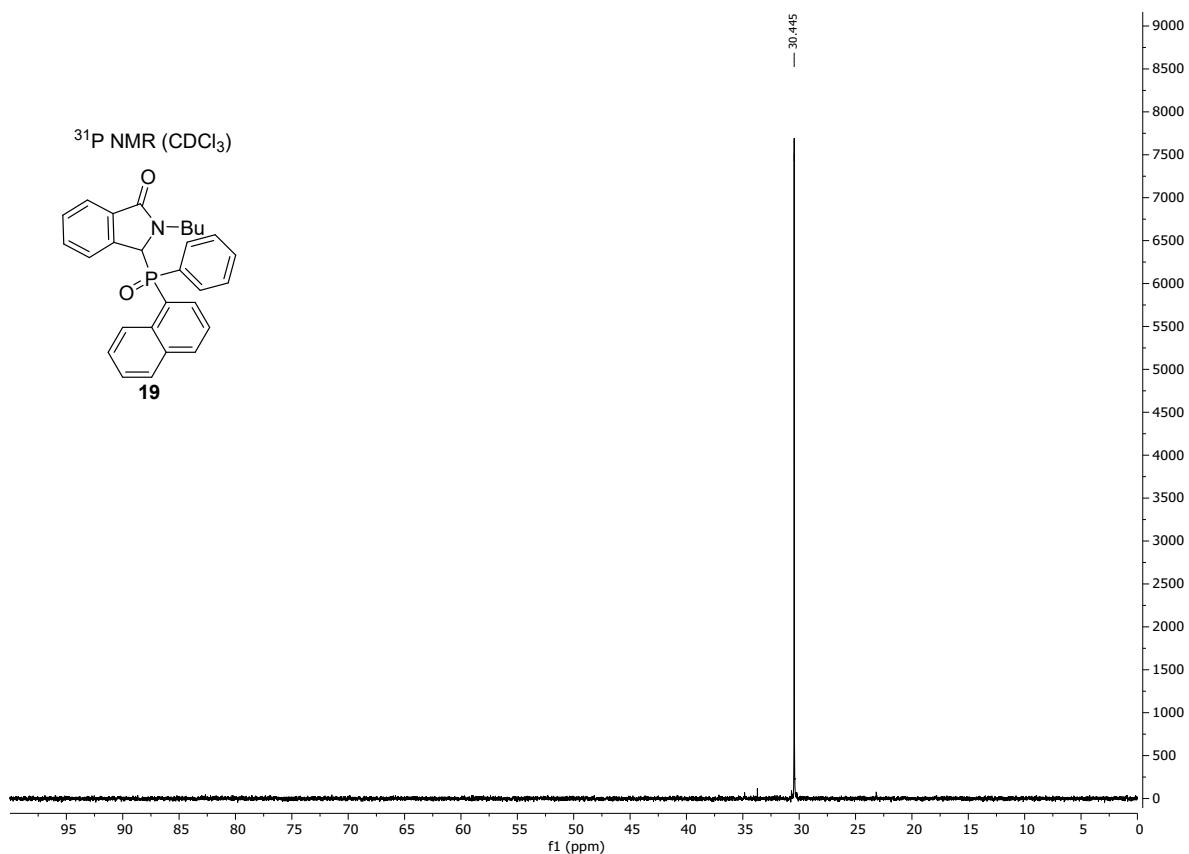
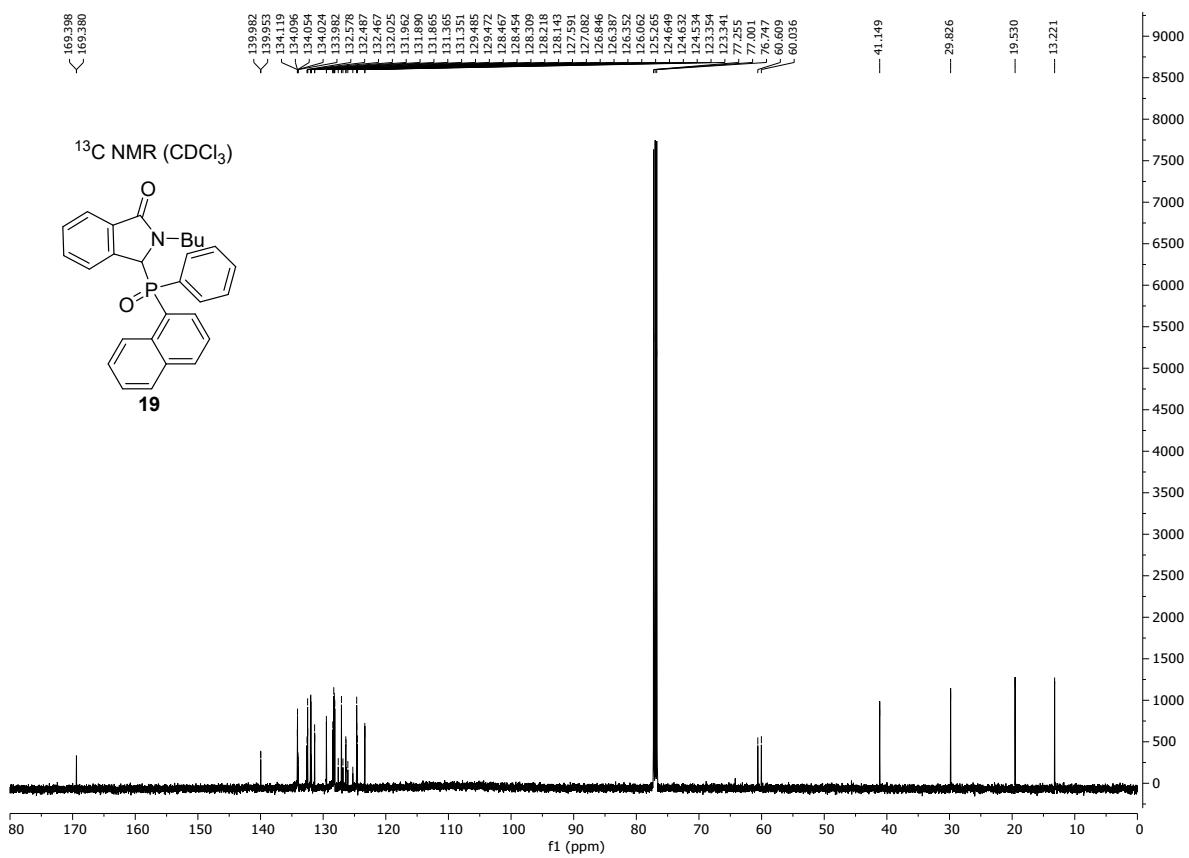


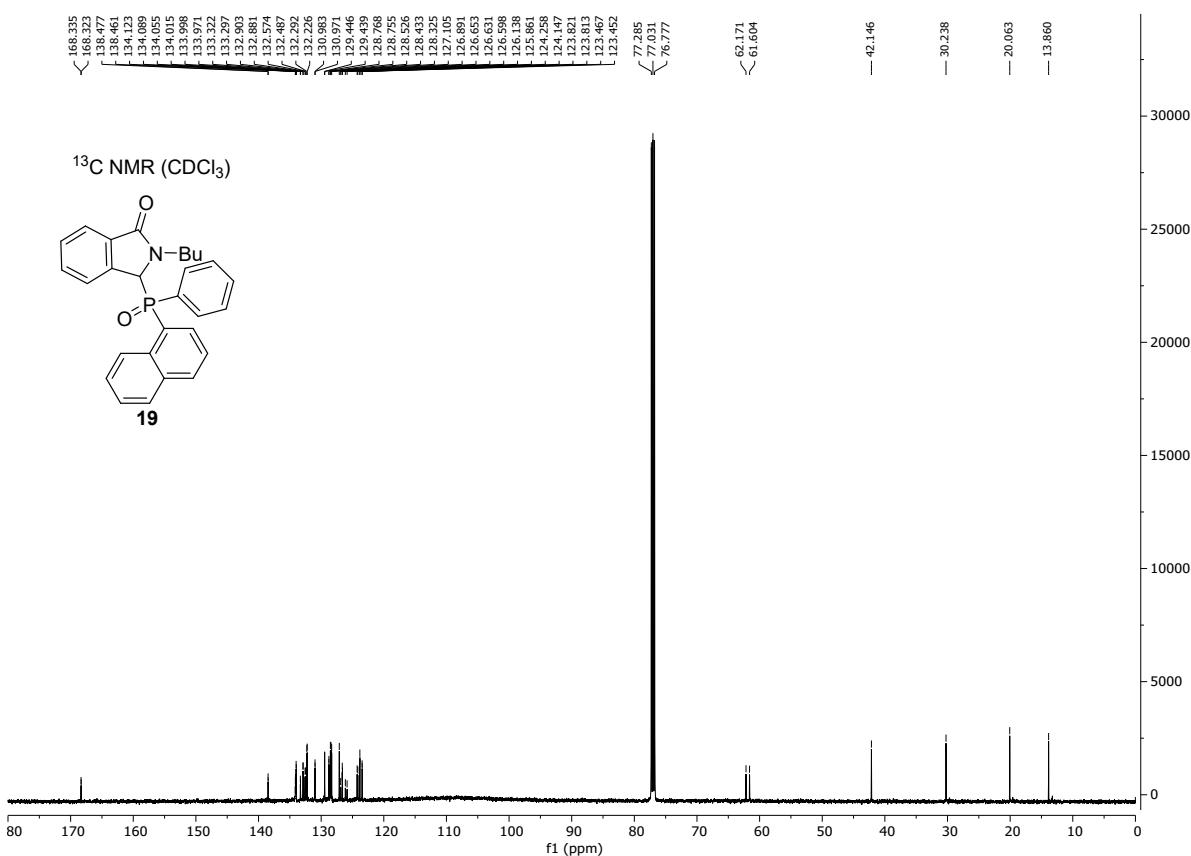
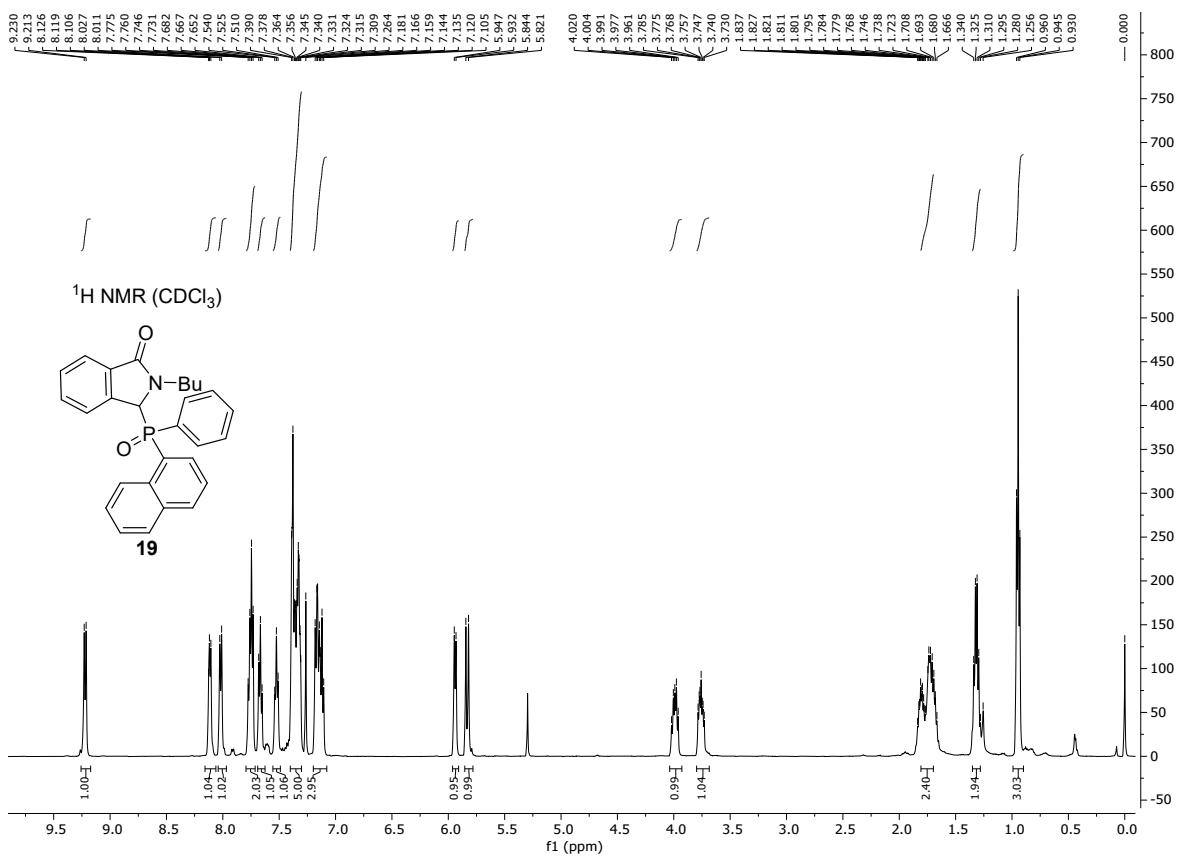


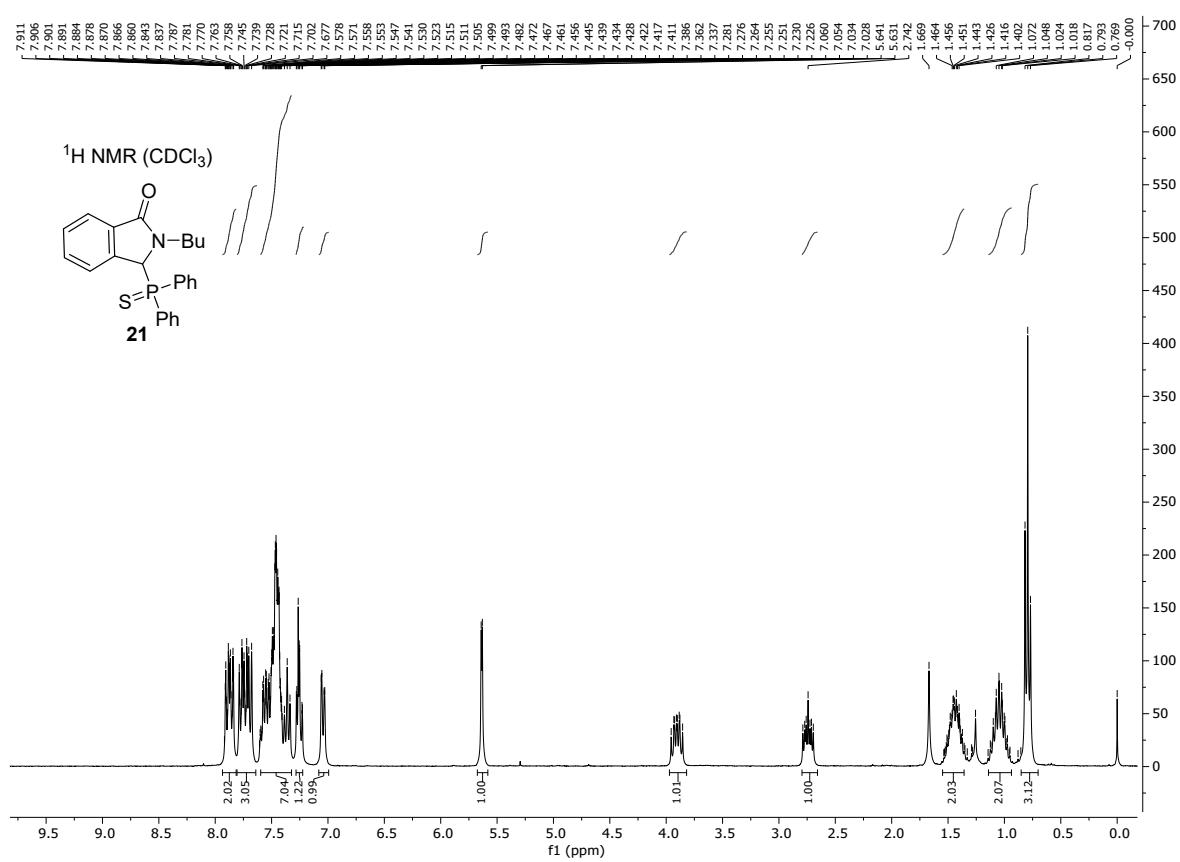
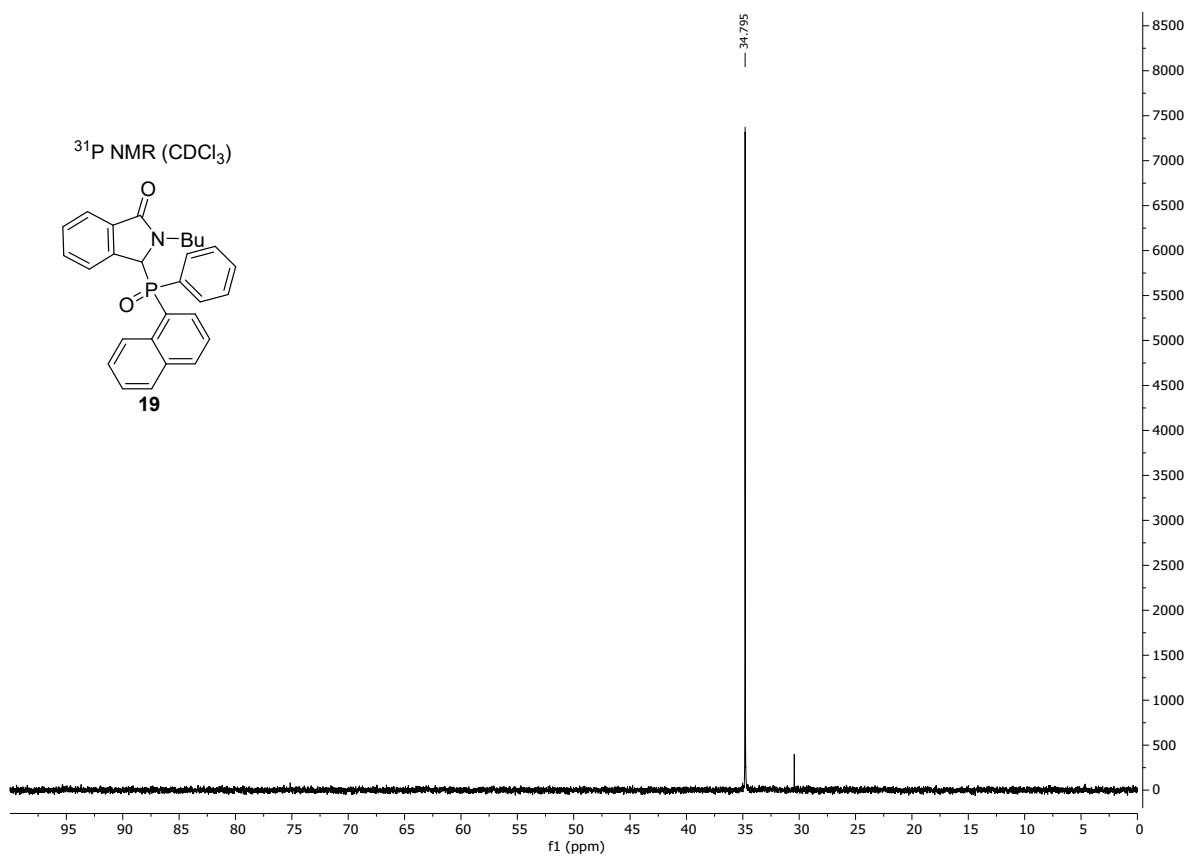


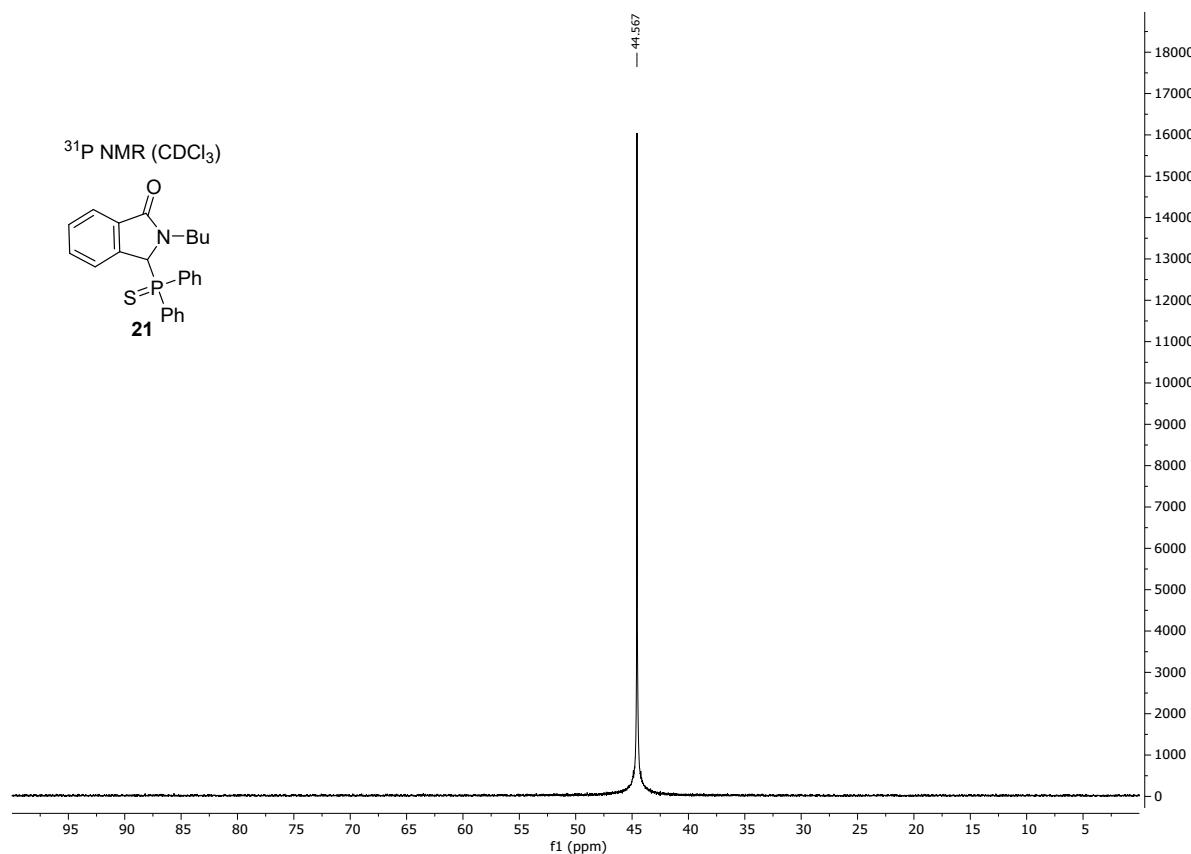
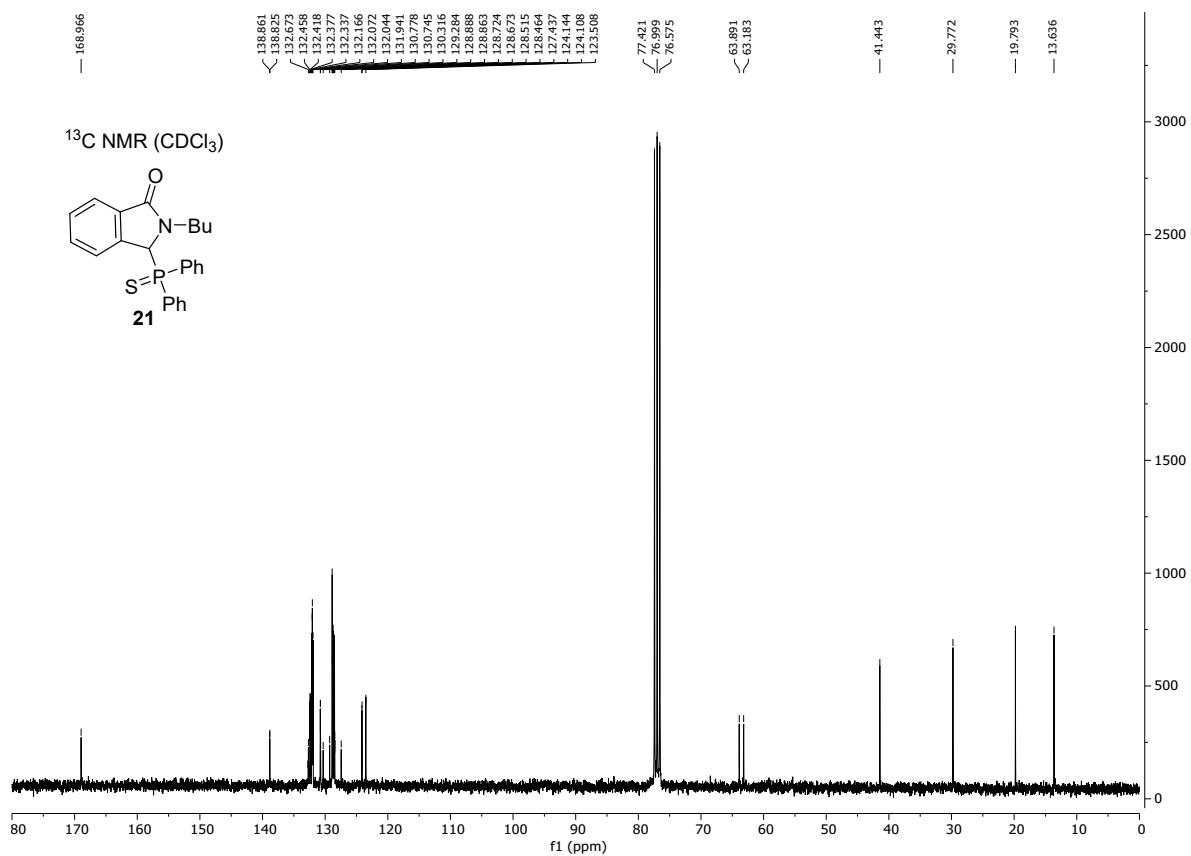


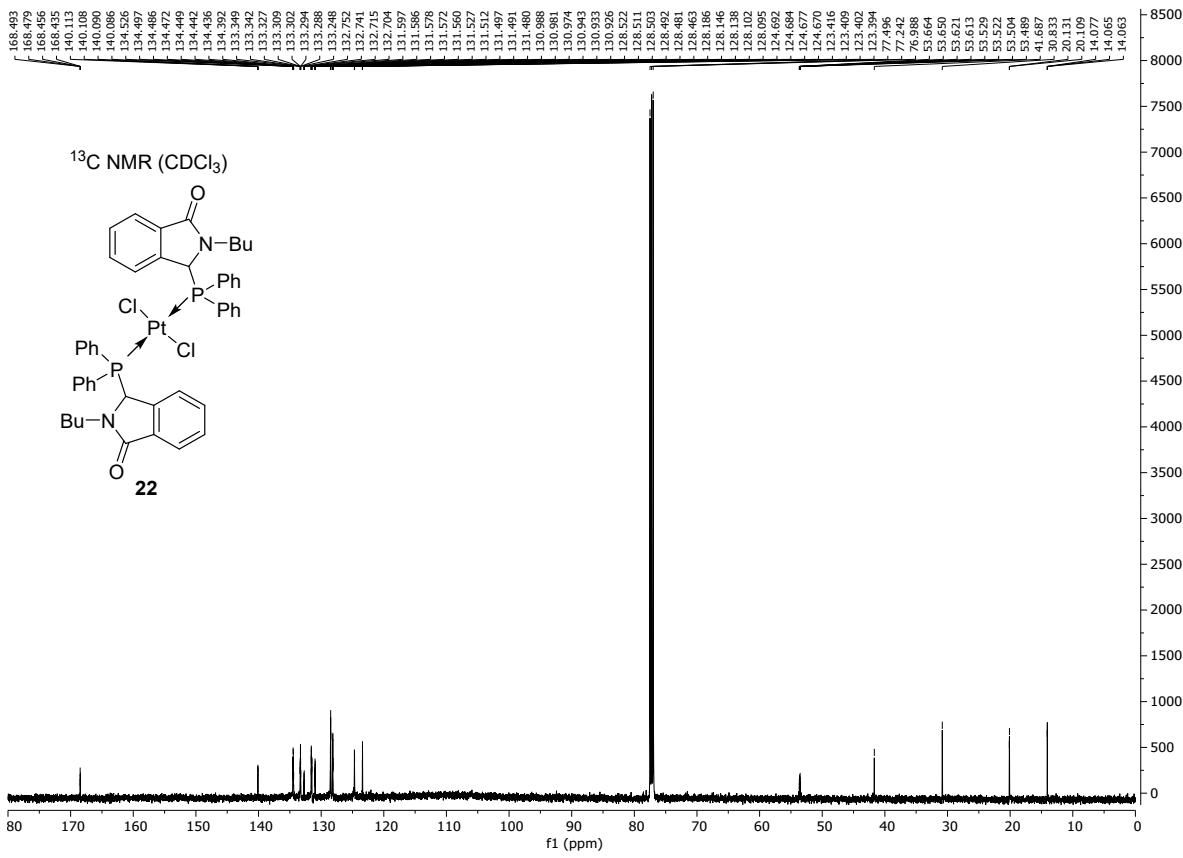
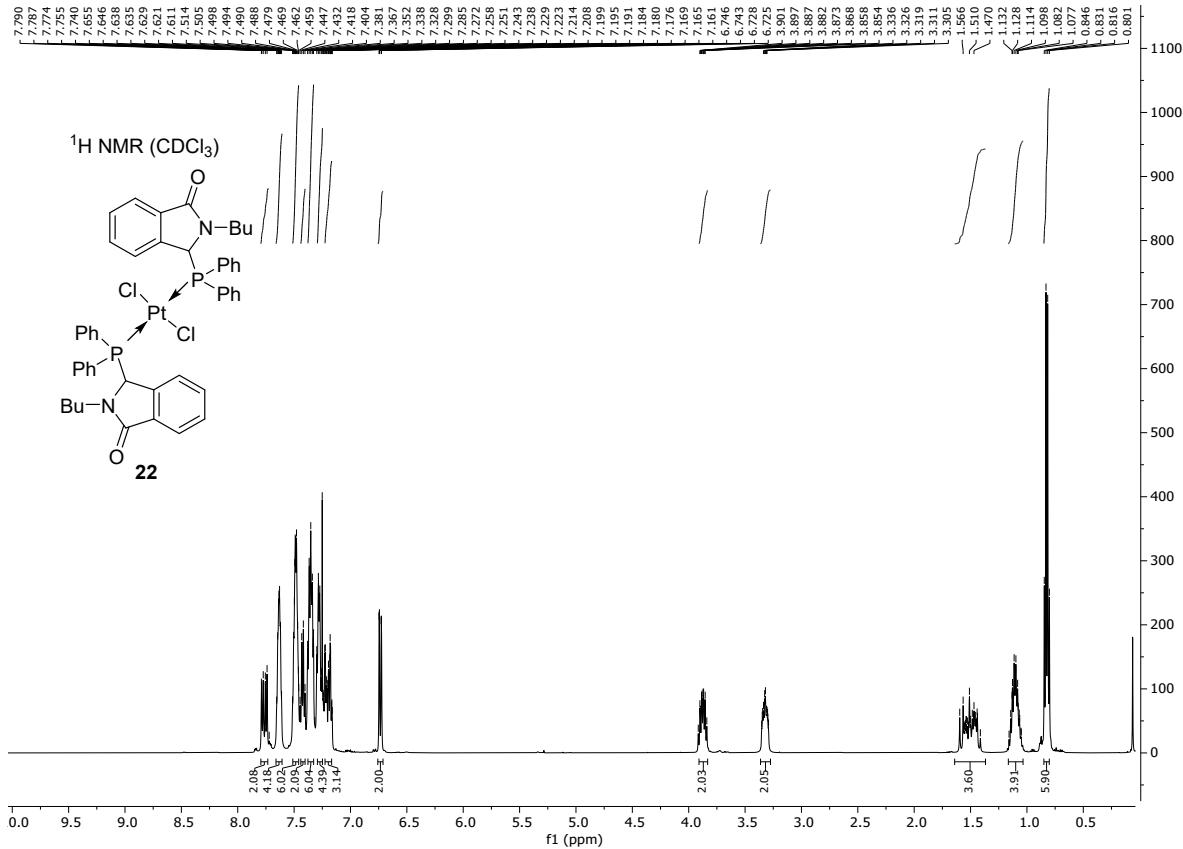


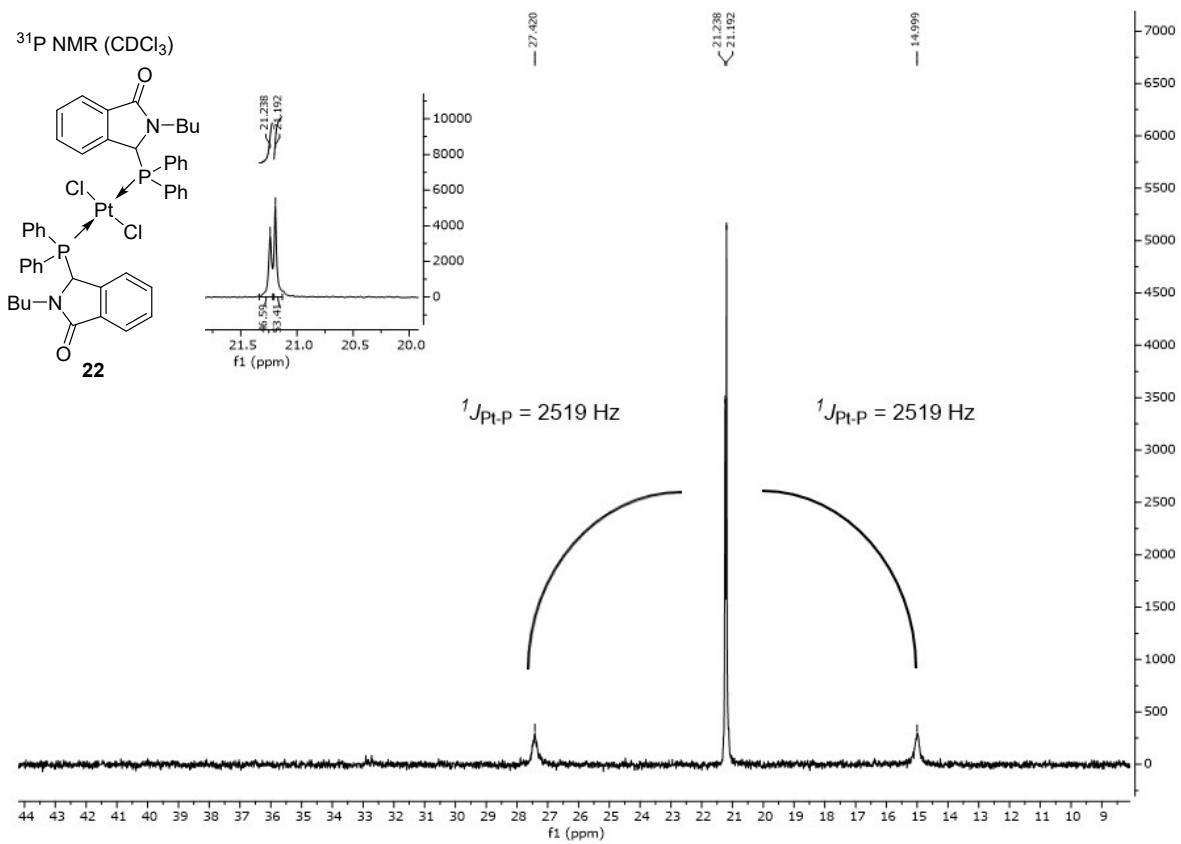
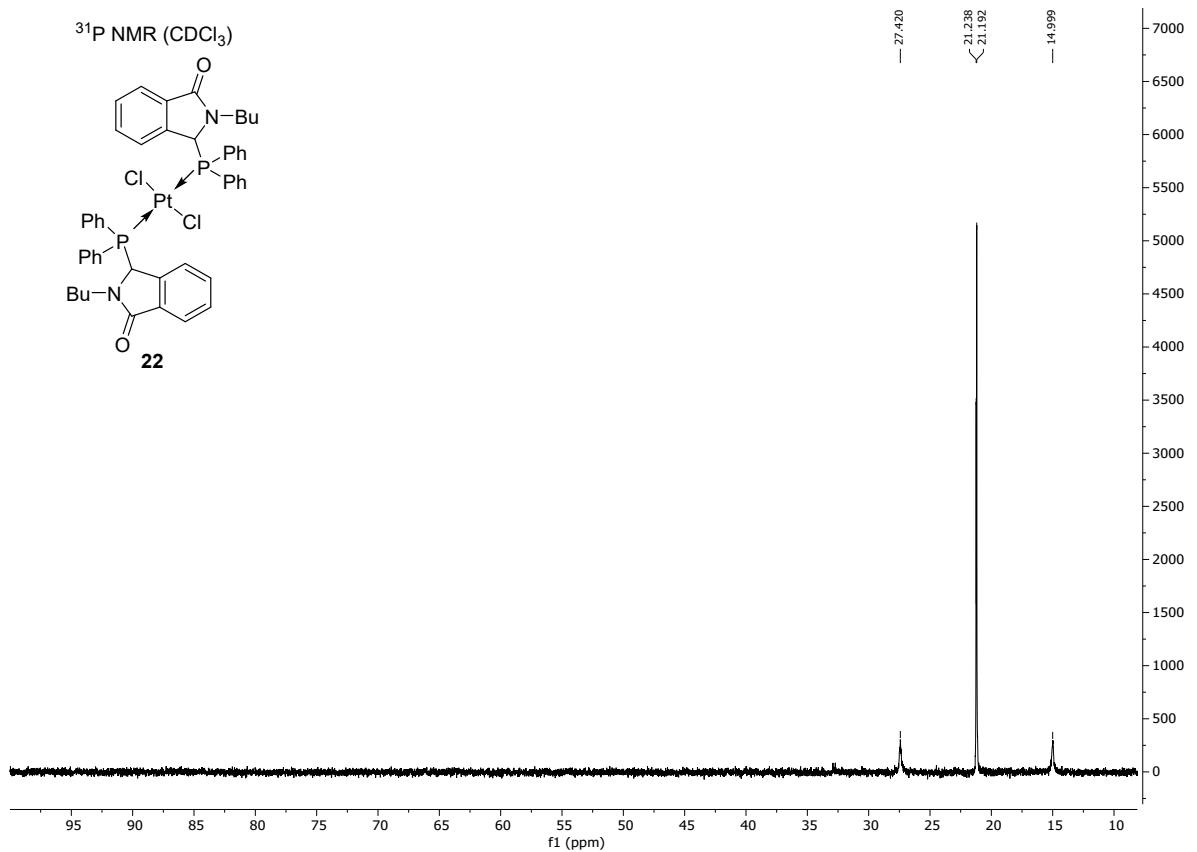












References

- [S1] *CrysAlisPro*, version 1.171.39.46e; Rigaku Oxford Diffraction: Yarnton, UK, 2018.
- [S2] G. M. Sheldrick, *Acta Cryst.* **2015**, *A71*, 3-8.
- [S3] G. M. Sheldrick, *Acta Cryst.* **2015**, *C71*, 3-8.
- [S4] O. V. Dolomanov, L. J. Bourhis, R. J. Gildea, J. A. K. Howard and H. Puschmann, *J. Appl. Cryst.* **2009**, *42*, 339-341.