Supporting Information

α-Amination of Keto-Nitrones via Multihetero-Cope Rearrangement Employing an Imidoyl Chloride Reagent

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Methods: General. Infrared (IR) spectra were obtained using a Jasco 460 Plus Fourier transform infrared spectrometer. Proton and carbon magnetic resonance spectra (¹H NMR and ¹³C NMR) were recorded on a Bruker model Avance 400 (¹H NMR at 400 MHz and ¹³C at 100 MHz) or a Bruker model Avance 600 (¹H NMR at 600 MHz and ¹³C NMR at 150 MHz) spectrometer with solvent resonance as the internal standard (¹H NMR: CDCl₃ at 7.26 ppm; ¹³C NMR: CDCl₃ at 77.0 ppm). ¹H NMR data are reported as follows: chemical shift, multiplicity (s = singlet, br s = broad singlet, d = doublet, t = triplet, br t = broad triplet, q = quartet, m = multiplet), coupling constants (Hz), and integration. Mass spectra were obtained using a Bruker BioTOF II spectrometer with electrospray ionization calibrated with CsOAc or an Agilent Technologies 6520, Accurate - Mass QTOF LCMS, 1200 series LC with dual spray ESI source. All samples were prepared in methanol. Analytical thin layer chromatography (TLC) was performed on Sorbent Technologies 0.20 mm Silica G TLC plates. Visualization was accomplished with UV light and/or aqueous ceric ammonium nitrate solution followed by heating. Purification of the reaction products was carried out by flash chromatography using Siliaflash-P60 silica gel (40-63µm) purchased from Silicycle. Unless otherwise noted, all reactions were carried out under an atmosphere of dry nitrogen in oven-dried glassware with magnetic stirring. Yield refers to isolated yield of analytically pure material unless otherwise noted. Yields are reported for a specific experiment and as a result may differ slightly from those found in the tables, which are averages of at least two experiments. Nitrone stereochemistry was assigned by ¹H NOESY analysis.

Materials: General. Dichloromethane was dried by passage through a column of neutral alumina under nitrogen prior to use. Triethylamine was freshly distilled from calcium hydride Benzyl-hydroxylamine,¹ *p*-methoxy-benzylhydroxylamine,² *p*-methylprior to use. benzylhydroxylamine.³ *p*-chloro-benzylhydroxylamine,³ and (S)- α -methylbenzylhydroxylamine⁴ were prepared according to the published procedures. Imidoyl chloride (1) was prepared via the literature method.⁵ Nitrone S11 was prepared via the published method.⁶ All other reagents were purchased from commercial sources and were used as received unless otherwise noted.

¹ Maskill, H.; Jencks, W. P. J. Am. Chem. Soc. 1987, 109, 2062.

² Nakano, Y.; Kato, Y.; Imai, K.; Ochiai, E.; Namekawa, J.; Ishizuka, S.; Takenouchi, K.; Tanatani, A.; Hashimoto, Y.; Nagasawa, K. *J. Med. Chem.* **2006**, *49*, 2398.

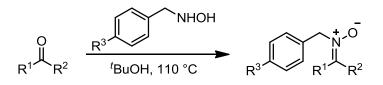
³ Gordon, I. M.; Maskill, H. J. Chem. Soc., Perkin Trans. 2 2001, 2059.

⁴ Breuning, M.; Hauser, T.; Tanzer, E.-M. Org. Lett., 2009, 11, 4032.

⁵ Osipov, S.; Artyushin, O.; Kolomiets, A.; Bruneau, C.; Dixneuf, P. Synlett, **2000**, 1031.

⁶ Pfeiffer, J. Y.; Beauchemin, A. M. J. Org. Chem., 2009, 74, 8381.

General Procedure A for Nitrone Synthesis:



An oven-dried 4-mL scintillation vial equipped with magnetic stir bar was charged with the appropriate ketone (2.0 mmol, 2.0 equiv) and hydroxylamine (1.0 mmol, 1 equiv). The vial was purged with nitrogen and tert-butanol (2.0 mL) was added. The septum was replaced with a Teflon-sealed cap and the reaction was stirred at 110 °C for 14 h. The reaction was cooled to 23 °C and volatiles were removed in vacuo. The crude product was purified via flash chromatography (SiO_2) to give the desired product.

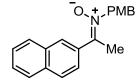
N-Cyclopentylidene-1-phenylmethanamine oxide (2): The title compound was prepared according to General Procedure A using cyclopentanone (1.00 mL, 11.3 mmol, 2.0 equiv) and benzylhydroxylamine (695 mg, 5.64 mmol, 1.0 equiv) in ^tBuOH (10 mL). Purification by flash chromatography (dichloromethane to 98:2 dichloromethane/methanol) furnished 2 as a tan solid (800 mg, 4.23 mmol, 75%). Analytical data: **mp** 64-67.5 °C; ¹**H NMR** (400 MHz, CDCl₃): δ 7.38 (d, *J* = 7.2 Hz, 2H,), 7.33-7.27 (m, 3H), 4.90 (s, 2H), 2.65 (br s, 2H), 2.50 (br s, 2H), 1.83-1.74 (m, 4H); ¹³C NMR (100 MHz, CDCl₃): δ 155.9, 133.2, 128.5, 128.0, 127.9, 65.0, 30.9, 30.8, 26.1, 24.3; HRMS (ESI⁺) Calcd. for C₁₂H₁₅NO+H, 190.1232; Found, 190.1229; **IR** (thin film, cm⁻¹) 3398, 2965, 2876, 1632, 1454, 1136, 960; **TLC** (95:5 dichloromethane/methanol): $R_f = 0.20$.

(Z)-1-Phenyl-N-(1-phenylethylidene)methanamine oxide (6): The title compound was prepared according to General Procedure A using acetophenone (0.234 mL, 2.0 mmol, 2.0 equiv) and benzyl-hydroxylamine (123 mg, 1.0 mmol, 1.0 equiv) in ^tBuOH (2 mL). Purification by flash chromatography (100:0 to 98:2 dichloromethane/methanol) furnished 6 as a pale yellow oil (111 mg, 0.493 mmol, 49%). Analytical data: ¹H NMR (400 MHz, CDCl₃): δ 7.42-7.39 (m, 3H), 7.31-7.28 (m, 5H), 7.21-7.19 (m, 2H), 4.97 (s, 2H), 2.45 (s, 3H); ¹³C NMR (100 MHz, CDCl₃): δ 147.6, 136.2, 134.2, 129.1, 128.8, 128.4, 128.1, 128.0, 127.5, 63.8, 20.6; HRMS (ESI⁺) Calcd. for C₁₅H₁₅NO+Na, 248.1052; Found, 248.1050; **IR** (thin film, cm⁻¹) 3223, 3060, 1579, 1495, 1453, 1265, 1178; **TLC** (95:5 dichloromethane/methanol): $R_f = 0.1$.

 $\bar{O}_{N'}$ PMB (Z)-1-(4-Methoxyphenyl)-N-(1-(4-methoxyphenyl)ethylidene) methanamine ovide (S1). The dial oxide (S1): The title compound was prepared according to General Procedure A using p-methoxyacetophenone (300 mg, 2.0 mmol, 2.0 equiv) and p-methoxybenzylhydroxylamine (153 mg, 1.0 mmol, 1.0 equiv) in ^tBuOH (2 mL).

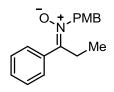
Purification by flash chromatography (100:0 to 98:2 dichloromethane/methanol) furnished S1 as

a yellow oil with a 6:1 diastereomeric ratio (37 mg, 0.130 mmol, 13%). Analytical data: ¹H **NMR** (400 MHz, CDCl₃): δ 7.23 (d, J = 8.4 Hz, 2H), 7.13 (d, J = 8.4 Hz, 2H), 6.92 (d, J = 8.8 Hz, 2H), 6.83 (d, J = 8.8 Hz, 2H), 4.89 (s, 2H), 3.82 (s, 3H), 3.76 (s, 3H), 2.39 (s, 3H); ¹³C NMR (100 MHz, CDCl₃): δ 160.0, 159.3, 147.0, 130.4, 130.1, 129.7, 129.0, 128.5, 126.5, 114.2, 114.1, 113.8, 113.1, 63.0, 55.3, 55.2, 55.2, 20.7, 19.2; **HRMS** (ESI⁺) Calcd. for C₁₇H₁₉NO₃+H, 286.1443; Found, 286.1444; **IR** (thin film, cm⁻¹) 3349, 2958, 2837, 2220, 1609, 1512, 1441, 1249, 1175; **TLC** (95:5 dichloromethane/methanol): R_f = 0.1.



(Z)-1-(4-Methoxyphenyl)-N-(1-(naphthalen-2-yl)ethylidene) methanamine oxide (S2): The title compound was prepared according to General Procedure A using 2-acetonaphthone (340 mg, 2.0 mmol, 2.0 equiv) and *p*-methoxy-benzylhydroxylamine (153 mg, 1.0 mmol, 1.0 equiv) in ^tBuOH (2 mL). Purification by flash chromatography (98:2

dichloromethane/methanol) furnished **S2** as a yellow oil (97.1 mg, 0.318 mmol, 32%). Analytical data: ¹**H NMR** (400 MHz, CDCl₃): δ 7.50-7.45 (m, 2H), 7.41-7.38 (m, 1H), 7.17-7.13 (m, 2H), 6.91 (d, *J* = 8.4 Hz, 1H), 6.83 (d, *J* = 8.8 Hz, 2H), 6.43 (d, *J* = 8.8 Hz, 2H), 4.53 (s, 2H), 3.37 (s, 3H), 2.09 (s, 3H); ¹³**C NMR** (100 MHz, CDCl₃): δ 159.5, 147.1, 133.6, 133.1, 132.8, 129.9, 128.9, 128.2, 127.8, 127.4, 127.3, 127.1, 126.5, 124.8, 113.9, 63.5, 55.3, 20.8; **HRMS (ESI**⁺) Calcd. for C₂₀H₁₉NO₂+Na, 328.1314; Found, 328.1324; **IR** (thin film, cm⁻¹) 3408, 3053, 2986, 1513, 1265, 1213, 1176; **TLC** (95:5 dichloromethane/methanol): R_f = 0.14.



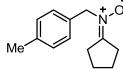
(Z)-1-(4-Methoxyphenyl)-N-(1-phenylpropylidene)methanamine oxide (S3): The title compound was prepared according to General Procedure A using propiophenone (0.26 mL, 2.0 mmol, 2.0 equiv) and *p*-methoxybenzylhydroxylamine (153 mg, 1.0 mmol, 1.0 equiv) in ^tBuOH (2 mL). Purification by flash chromatography (98:2 dichloromethane/methanol) furnished S3 as a pale yellow oil (83.2 mg, 0.309 mmol, 31%). Analytical

data: ¹H NMR (400 MHz, CDCl₃): δ 7.44-7.42 (m, 3H), 7.20-7.15 (m, 4H), 6.82 (d, J = 8.8 Hz, 2H), 4.80 (s, 2H), 3.78 (s, 3H), 2.88 (q, J = 7.2, 7.6 Hz, 2H), 1.02 (t, J = 7.4 Hz, 3H); ¹³C NMR (100 MHz, CDCl₃): δ 159.4, 151.6, 135.1, 129.9, 129.1, 128.9, 128.0, 126.4, 113.8, 63.8, 55.2, 26.2, 9.3; HRMS (ESI⁺) Calcd. for C₁₇H₁₉NO₂+H, 270.1495; Found, 270.1497; IR (thin film, cm⁻¹) 3409, 3053, 2985, 1612, 1493, 1265, 1177; TLC (95:5 dichloromethane/methanol): R_f = 0.16.

N-Cyclopentylidene-1-(4-methoxyphenyl)methanamine oxide (S4): The title compound was prepared according to General Procedure A using cyclopentanone (0.177 mL, 2.0 mmol, 2.0 equiv) and p-methoxy-benzylhydroxylamine (153 mg, 1.00 mmol, 1.0 equiv) in ^tBuOH (2 mL). Purification by flash chromatography (100:0 to 95:5 dichloromethane/methanol) furnished S4 as a brown oil (192 mg,

 $(100.0 \text{ to } 95.5 \text{ diction of neural e/methanol) furthistical 34 as a brown of (192 mg, 0.876 mmol, 88%). Analytical data: ¹H NMR (400 MHz, CDCl₃): <math>\delta$ 7.32 (d, J = 8.8 Hz, 2H), 6.82 (d, J = 8.8 Hz, 2H), 4.82 (s, 2H), 3.73 (s, 2H), 2.61 (t, J = 7.2 Hz, 2H), 2.50 (t, J = 7.2 Hz, 2H), 1.81-1.72 (m, 4H); ¹³C NMR (100 MHz, CDCl₃): δ 159.4, 155.0, 129.5, 125.5, 113.9, 64.5,

55.1, 30.9, 26.2, 24.3; **HRMS** (**ESI**⁺) Calcd. for $C_{13}H_{17}NO_2$ +H, 220.1337; Found, 220.1338; **IR** (thin film, cm⁻¹) 3388, 2963, 1612, 1513, 1250, 1179, 1133, 1029; **TLC** (95:5 dichloromethane/methanol): $R_f = 0.15$.



N-Cyclopentylidene-1-(p-tolyl)methanamine oxide (S5): The title compound was prepared according to General Procedure A using cyclopentanone (0.18 mL, 2.0 mmol, 2.0 equiv) and *p*-methylbenzylhydroxylamine (137 mg, 1.0 mmol, 1.0 equiv) in ^{*t*}BuOH (2 mL). Purification by flash chromatography (98:2 to 95:5

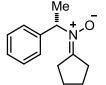
dichloromethane/methanol) furnished **S5** as an off-white semi-solid (160 mg, 0.787 mmol, 77%). Analytical data: ¹**H NMR** (400 MHz, CDCl₃): δ 7.29 (d, *J* = 8.0 Hz, 2H), 7.13 (d, *J* = 9.6 Hz, 2H), 4.88 (s, 2H), 2.65 (t, *J* = 7.2, 2H), 2.51 (t, *J* = 6.8 Hz, 2H), 2.31 (s, 3H), 1.85-1.75 (m, 4H); ¹³**C NMR** (100 MHz, CDCl₃): δ 155.2, 137.8, 130.3, 129.2, 128.0, 64.9, 30.9, 30.9, 26.2, 24.3, 20.9; **HRMS (ESI**⁺) Calcd. for C₁₃H₁₇NO+Na, 226.1208; Found, 226.1208; **IR** (thin film, cm⁻¹) 3388, 2964, 2875, 2207, 1627, 1514, 1452, 1142; **TLC** (95:5 dichloromethane/methanol): R_f = 0.4.

1-(4-Chlorophenyl)-N-cyclopentylidenemethanamine oxide (S6): The title compound was prepared according to General Procedure A using cyclopentanone (0.18 mL, 2.0 mmol, 2.0 equiv) and pchlorobenzylhydroxylamine (157 mg, 1.0 mmol, 1.0 equiv) in ^tBuOH (2 Purification flash chromatography mL). by (98:2)dichloromethane/methanol) furnished S6 as an off-white solid (175 mg, 0.784 mmol, 78%). Analytical data: **mp** 104-107 °C; ¹**H NMR** (400 MHz, CDCl₃): δ 7.39 (d, J = 8.4 Hz, 2H), 7.34 (d, J = 8.4 Hz, 2H), 4.92 (s, 2H), 2.69 (t, J = 7.2 Hz, 2H), 2.54 (t, J = 7.0 Hz, 2H), 1.90-1.80 (m, 4H); ¹³C NMR (100 MHz, CDCl₃): δ 155.8, 134.2, 131.9, 129.6, 128.9, 64.3, 31.1, 31.1, 26.3, 24.5; **HRMS** (**ESI**⁺) Calcd. for C₁₂H₁₄ClNO+H, 224.0841; Found, 224.0843; **IR** (thin film, cm⁻¹) 3388, 2964, 2875, 2207, 1627, 1514, 1452, 1142; TLC (95:5 dichloromethane/methanol): R_f = 0.12.

- *N*-Cyclohexylidene-1-(4-methoxyphenyl)methanamine oxide (S7): The title compound was prepared according to General Procedure A using cyclohexanone (0.2 mL, 2.00 mmol, 2.0 equiv) and *p*-methoxybenzylhydroxylamine (153 mg, 1.00 mmol, 1.0 equiv) in ^tBuOH (2 mL). Purification by flash chromatography (100:0 to 95:5 dichloromethane/methanol) furnished S7 as a yellow oil (167 mg, 0.716 mmol,

71%). Analytical data: ¹**H NMR** (400 MHz, CDCl₃): δ 7.30 (d, J = 8.8 Hz, 2H), 6.83 (d, J = 8.4 Hz, 2H), 4.98 (s, 2H), 3.74 (s, 3H), 2.71 (t, J = 6.4, 2H), 2.46 (t, J = 6.0 Hz, 2H), 1.65-1.59 (m, 2H), 1.56-1.48 (m, 4H); ¹³**C NMR** (100 MHz, CDCl₃): δ 159.3, 149.3, 129.0, 126.2, 114.0, 63.0, 55.1, 29.9, 26.8, 25.5, 24.5; **HRMS** (**ESI**⁺) Calcd. for C₁₄H₁₉NO+H, 234.1494; Found, 234.1484; **IR** (thin film, cm⁻¹) 3400, 2937, 2860, 1612, 1513, 1249, 1032, 734; **TLC** (95:5 dichloromethane/methanol): $\mathbf{R}_{\rm f} = 0.2$.

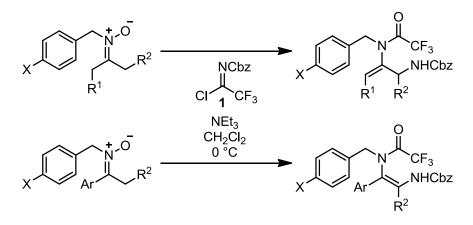
N-(4-(Tert-butyl)cyclohexylidene)-1-(4-methoxyphenyl)methanamine oxide ±,PMB (S8): The title compound was prepared according to General Procedure A using 4butylcyclohexanone (308)mg, 2.0 mmol. 2.0equiv) **p**and methoxybenzylhydroxylamine (153 mg, 1.0 mmol, 1.0 equiv) in ^tBuOH (2 mL). Purification by flash chromatography (100:0 to 98:2 dichloromethane/methanol) furnished **S8** as a pale vellow solid (220 mg, 0.761 mmol, 76%). Analytical data: ^tBu **mp** 74-77.5 °C; ¹**H NMR** (400 MHz, CDCl₃): δ 7.34 (d, J = 8.8 Hz, 2H), 6.88 (d, J = 8.8 Hz, 2H), 5.03 (d, J = 14.0 Hz, 1H), 4.99 (d, J = 14.0 Hz, 1H), 3.79 (s, 3H), 3.60-3.56 (m, 1H), 2.88-2.84 (m, 1H), 2.01-1.86 (m, 3H), 1.25-1.15 (m, 3H), 0.812 (s, 9H); ¹³C NMR (100 MHz, CDCl₃): 8 159.3, 149.3, 129.1, 126.1, 114.0, 63.0, 55.1, 46.1, 32.3, 29.8, 27.3, 27.0, 26.2, 25.0; **HRMS** (**ESI**⁺) Calcd. for C₁₈H₂₇NO₂+H, 290.2120; Found, 290.2121; **IR** (thin film, cm⁻¹) 3409, 3053, 2985, 1612, 1493, 1265, 1177; **TLC** (95:5 dichloromethane/methanol): R_f = 0.16.



(S)-N-Cyclopentylidene-1-phenylethanamine oxide (S9): The title compound was prepared according to General Procedure A using cyclopentanone (0.18 mL, 2.0 mmol, 2.0 equiv) and (S)- α -methyl-benzyl-hydroxylamine⁴ (137 mg, 1.0 mmol, 1.0 equiv) in ^tBuOH (2 mL). Purification by flash chromatography (98:2 dichloromethane/methanol) furnished **S9** as an off-white solid (177.0 mg,

0.871 mmol, 87%). Analytical data: **mp** 76-79 °C; ¹**H NMR** (400 MHz, CDCl₃): δ 7.47 (d, J = 7.6 Hz, 2H), 7.35-7.28 (m, 3H), 5.15 (q, J = 6.8 Hz, 1H), 2.75-2.49 (m, 4H), 1.88-1.78 (m, 4H), 1.75 (d, J = 6.8 Hz, 3H); ¹³**C NMR** (100 MHz, CDCl₃): δ 154.2, 139.1, 128.6, 128.1, 127.0, 67.7, 31.2, 30.6, 26.3, 24.3, 19.3; **HRMS** (**ESI**⁺) Calcd. for C₁₃H₁₇NO+Na, 226.1208; Found, 226.1214; **IR** (thin film, cm⁻¹) 3399, 2967, 2876, 2190, 1622, 1454, 1255, 1172, 1141, 1055; **TLC** (95:5 dichloromethane/methanol): R_f = 0.13.

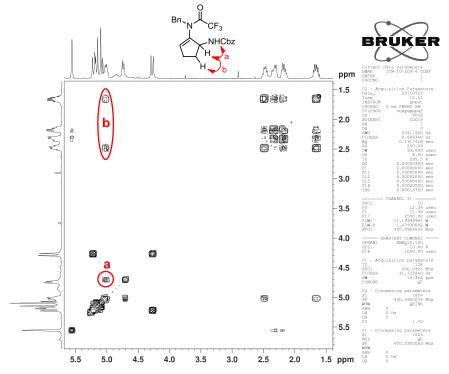
General Procedure B for [3,3]-Rearrangement:



An oven-dried 4-mL scintillation vial equipped with magnetic stir bar was charged with imidoyl chloride **1** (0.26 g, 0.1 mmol, 1.0 equiv) and purged with nitrogen. Dichloromethane (0.5 mL) was added and the solution was cooled to 0 °C in an ice/water bath. Triethylamine (0.028 mL, 0.2 mmol, 2.0 equiv) was then added, followed by a cooled dichloromethane solution (0.5 mL) of nitrone **2**, **6**, **S1-S11** (0.1 mmol, 1.0 equiv). The reaction was stirred at 0 °C until judged complete by TLC analysis, generally 30 min. Upon completion, volatiles were removed *in vacuo*. The crude product was purified via flash chromatography (SiO₂) to give the desired product.

Benzyl (2-(N-benzyl-2,2,2-trifluoroacetamido)cyclopent-2-en-1-yl) carbamate (3): The title compound was prepared according to General Procedure B using nitrone 2 (57 mg, 0.305 mmol, 1.0 equiv), imidoyl chloride 1 (81 mg, 0.305 mmol, 1.0 equiv) and triethylamine (85 μL, 0.61 mmol, 2.0 equiv) in CH₂Cl₂ (2 mL). Purification by flash chromatography (90:10 petroleum ether/acetone) furnished 3 as a yellow oil (82 mg, 0.196 mmol, 1.9 mmol, 1.9 mmol).

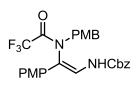
65%). Analytical data: ¹**H NMR** (400 MHz, CDCl₃): δ 7.38-7.25 (m, 10H), 5.56 (s, 1H), 5.21 (d, J = 14.4 Hz, 1H), 5.16 (d, J = 12.4 Hz, 1H), 5.08 (d, J = 12.0 Hz, 1H), 5.05-4.99 (m, 1H), 4.74 (d, J = 9.6 Hz, 1H), 4.28 (d, J = 14.8 Hz, 1H), 2.49-2.44 (m, 1H), 2.36-2.30 (m, 1H), 2.21-2.15 (m, 1H), 1.70-1.63 (m, 1H); ¹³C NMR (100 MHz, CDCl₃): δ 157.1, 156.8, 155.9, 137.8, 136.1, 135.1, 132.9, 128.6, 128.6, 128.2, 128.1, 127.9, 67.0, 54.0, 49.6, 30.9, 27.1; **HRMS** (**ESI**⁺) Calcd. for C₂₂H₂₁F₃N₂O₃+Na, 441.1402; Found, 441.1402; **IR** (thin film, cm⁻¹) 3428, 3352, 3064, 2950, 1693, 1512, 1208, 1157, 1051; **TLC** (60:40 petroleum ether/acetone): R_f = 0.60.



¹H COSY analysis of 3 supporting acyl transfer:

 $\begin{array}{ll} \textbf{(Z)-Benzyl} & \textbf{(2-(N-benzyl-2,2,2-trifluoroacetamido)-2-phenylvinyl)} \\ \textbf{(Z)-Benzyl} & \textbf{(2-(N-benzyl-2,2,2-trifluoroacetamido)-2-phenylvinyl)} \\ \textbf{(arbamate (7): The title compound was prepared according to General Procedure B using nitrone$ **6**(22 mg, 0.10 mmol, 1.0 equiv), imidoyl chloride**1** $(26.5 mg, 0.1 mmol, 1.0 equiv) and triethylamine (28 <math>\mu$ L, 0.20 mmol, 2.0 equiv) in CH₂Cl₂ (1 mL). Purification by flash chromatography (90:10

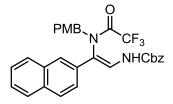
petroleum ether/acetone) furnished **7** as a colorless oil (28 mg, 0.062 mmol, 61%). Analytical data: ¹H NMR (400 MHz, CDCl₃): δ 7.40-7.15 (m, 15H), 5.70 (d, *J* = 10.8 Hz, 1H), 5.60 (d, *J* = 13.2 Hz, 1H), 5.10 (d, *J* = 12.4 Hz, 1H), 4.92 (d, *J* = 12.0 Hz, 1H), 3.76 (d, *J* = 13.6 Hz, 1H); ¹³C NMR (100 MHz, CDCl₃): δ 158.7, 158.3, 152.6, 135.6, 135.3, 133.6, 129.5, 129.1, 129.0, 128.8, 128.4, 128.4, 128.0, 127.7, 123.8, 123.5, 117.7, 67.5, 51.6; HRMS (ESI⁺) Calcd. for C₂₅H₂₁F₃N₂O₃+H, 455.1582; Found, 455.1585; IR (thin film, cm⁻¹) 3409, 3053,1691, 1665, 1482, 1265, 1211, 1174; TLC (75:25 hexanes/ethyl acetate): R_f = 0.30.



(Z)-Benzyl (2-(4-methoxyphenyl)-2-(2,2,2-trifluoro-N-(4-methoxy benzyl)acetamido)vinyl) carbamate (10): The title compound was prepared according to General Procedure B using nitrone S1 (30 mg, 0.105 mmol, 1.0 equiv), imidoyl chloride 1 (28 mg, 0.105 mmol, 1.0 equiv) and triethylamine (30 μ L, 0.21 mmol, 2.0 equiv) in CH₂Cl₂ (1 mL).

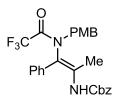
Purification by flash chromatography (90:10 petroleum ether/acetone) furnished **10** as a pale yellow oil (37 mg, 0.072 mmol, 69%). Analytical data: ¹H NMR (600 MHz, CDCl₃): δ 7.39-7.22 (m, 9H), 6.92 (d, J = 9.0 Hz, 2H), 6.82 (d, J = 8.4 Hz, 2H), 5.72 (d, J = 11.4 Hz, 1H), 5.50 (d, J = 11.4 Hz, 1H), 5.50 (d, J = 8.4 Hz, 2H), 5.72 (d, J = 11.4 Hz, 1H), 5.50 (d, J = 11.4 H

13.2 Hz, 1H), 5.06 (d, J = 12.6 Hz, 1H), 4.97 (d, J = 12.6 Hz, 1H), 3.83 (s, 3H), 3.72 (d, J = 13.8 Hz, 1H), 3.67 (s, 3H); ¹³C NMR (150 MHz, CDCl₃): δ 160.0, 159.4, 152.8, 135.4, 130.9, 128.8, 128.6, 128.3, 127.7, 126.0, 125.4, 121.8, 117.9, 114.5, 114.5, 67.5, 55.3, 55.2, 50.9; **HRMS** (ESI⁺) Calcd. for C₂₇H₂₅F₃N₂O₅+H, 515.1794; Found, 515.1794; **IR** (thin film, cm⁻¹) 3403, 3057, 2959, 1693, 1611, 1483, 1215, 1033; **TLC** (70:30 petroleum ether/acetone): R_f = 0.60.



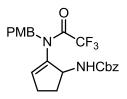
(Z)-Benzyl (2-(naphthalen-2-yl)-2-(2,2,2-trifluoro-*N*-(4-methoxy benzyl)acetamido)vinyl) carbamate (11): The title compound was prepared according to General Procedure B using nitrone S2 (30.5 mg, 0.10 mmol, 1.0 equiv), imidoyl chloride 1 (26.5 mg, 0.1 mmol, 1.0 equiv) and triethylamine (27.9 μ L, 0.20 mmol, 2.0 equiv) in CH₂Cl₂ (1 mL). Purification by flash chromatography (90:10

hexanes/ethyl acetate) furnished **11** as an off-white solid (21.7 mg, 0.041 mmol, 51%). Analytical data: **mp** 50-51 °C; ¹**H NMR** (400 MHz, CDCl₃): δ 7.84 (q, *J* = 8.4, 6.8 Hz, 3H), 7.59 (br s, 1H), 7.52-7.50 (m, 4H), 7.40-7.27 (m, 3H), 6.83 (d, *J* = 8.8, 2H), 5.87 (d, *J* = 11.2 Hz, 1H), 5.61 (d, *J* = 13.6 Hz, 1H), 5.10 (d, *J* = 12.4 Hz, 1H), 5.01 (d, *J* = 12.0 Hz, 1H), 3.84 (d, *J* = 13.6 Hz, 1H), 3.67 (s, 3H); ¹³C **NMR** (100 MHz, CDCl₃): δ 160.1, 152.8, 135.2, 133.4, 132.7, 131.0, 130.9, 128.9, 128.6, 128.4, 128.1, 127.8, 127.6, 126.8, 126.3, 123.9, 122.7, 121.6, 118.0, 117.0, 115.1, 114.6, 67.7, 55.2, 51.3; **HRMS** (**ESI**⁺) Calcd. for C₃₀H₂₅F₃N₂O₄+H, 535.1845; Found, 535.1838; **IR** (thin film, cm⁻¹) 3408, 3054, 2986, 2305, 1692, 1659, 1512, 1484, 1265, 1213; **TLC** (70:30 hexanes/ethyl acetate): R_f = 0.51.

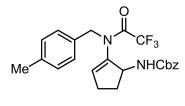


(*E*)-Benzyl (1-phenyl-1-(2,2,2-trifluoro-*N*-(4-methoxybenzyl)acetamido) prop-1-en-2-yl) carbamate (12): The title compound was prepared according to General Procedure B using nitrone S3 (27 mg, 0.10 mmol, 1.0 equiv), imidoyl chloride 1 (27 mg, 0.10 mmol, 1.0 equiv) and triethylamine (28 μ L, 0.20 mmol, 2.0 equiv) in CH₂Cl₂ (1 mL). Purification by flash chromatography (90:10 hexanes/ethyl acetate) furnished 12 as a pale yellow

oil (31.1 mg, 0.062 mmol, 62%) as a 5.5:1 mixture of inseparable diastereomers. Analytical data: ¹H NMR (400 MHz, CDCl₃): (Major diastereomer) δ 7.44-7.28 (m, 10H), 7.24 (d, *J* = 7.2 Hz, 2H), 6.84 (d, *J* = 8.8 Hz, 2H), 5.72 (s, 1H), 5.30 (d, *J* = 13.6 Hz, 1H), 4.98 (d, *J* = 12.4 Hz, 1H), 4.92 (d, *J* = 12.4 Hz, 1H), 3.69 (s, 3H), 3.51 (d, *J* = 13.6 Hz, 1H), 2.31 (s, 3H); ¹³C NMR (100 MHz, CDCl₃): δ 160.0, 152.5, 135.8, 135.5, 133.9, 130.8, 129.8, 129.7, 128.7, 128.6, 128.5, 128.3, 128.1, 127.9, 127.7, 127.5, 119.0, 114.7, 114.1, 66.7, 55.2, 50.1, 16.5; HRMS (ESI⁺) Calcd. for C₂₇H₂₅F₃N₂O₄+Na, 521.1665; Found, 521.1664; IR (thin film, cm⁻¹) 3387, 2593, 1690, 1641, 1512, 1482, 1250, 1167, 1110; TLC (70:30 hexane/ethyl acetate): R_f = 0.58.

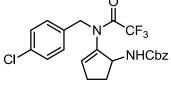


Benzyl (2-(2,2,2-trifluoro-*N*-(4-methoxybenzyl)acetamido) cyclopent-2en-1-yl)carbamate (13): The title compound was prepared according to General Procedure B using nitrone S4 (38 mg, 0.173 mmol, 1.0 equiv), imidoyl chloride 1 (46 mg, 0.173 mmol, 1.0 equiv) and triethylamine (48 μ L, 0.346 mmol, 2.0 equiv) in CH₂Cl₂ (2 mL). Purification by flash chromatography (90:10 petroleum ether/acetone) furnished **13** as a colorless oil (52 mg, 0.116 mmol, 67%). Analytical data: ¹H NMR (600 MHz, CDCl₃): δ 7.38-7.34 (m, 5H), 7.18 (d, *J* = 8.4 Hz, 2H), 6.83 (d, *J* = 9.0 Hz, 2H), 5.52 (s, 1H), 5.16 (d, *J* = 12.0 Hz, 1H), 5.07 (d, *J* = 12.0 Hz, 1H), 5.02-4.98 (m, 1H), 4.68 (d, *J* = 9.6 Hz, 1H), 4.17 (d, *J* = 14.4 Hz, 1H), 3.78 (s, 3H), 2.50-2.47 (m, 1H), 2.35-2.30 (m, 1H), 2.20-2.17 (m, 1H), 1.66-1.62 (m, 1H); ¹³C NMR (150 MHz, CDCl₃): δ 159.3, 155.9, 137.7, 136.1, 132.9, 130.1, 128.6, 128.5, 128.4, 128.1, 127.2, 113.9, 68.1, 67.1, 55.2, 54.0, 49.1, 30.9, 27.1; HRMS (ESI⁺) Calcd. for C₂₃H₂₃F₃N₂O₄+Na, 471.1508; Found, 471.1508; **IR** (thin film, cm⁻¹) 3426, 3054, 1691, 1513, 1265, 1207, 1175; **TLC** (70:30 petroleum ether/acetone): R_f = 0.80.



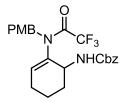
Benzyl (2-(2,2,2-trifluoro-*N*-(4-methylbenzyl)acetamido) cyclopent-2-en-1-yl)carbamate (14): The title compound was prepared according to General Procedure B using nitrone S5 (20 mg, 0.10 mmol, 1.0 equiv), imidoyl chloride 1 (27 mg, 0.10 mmol, 1.0 equiv) and triethylamine (28 μ L, 0.20 mmol, 2.0 equiv) in CH₂Cl₂ (1 mL). Purification by flash chromatography (90:10

petroleum ether/acetone) furnished **14** as a colorless oil (34 mg, 0.079 mmol, 79%). Analytical data: ¹**H NMR** (400 MHz, CDCl₃): δ 7.40-7.33 (m, 5H), 7.15-7.10 (m, 4H), 5.55 (s, 1H), 5.16 (d, J = 12.4 Hz, 2H), 5.08 (d, J = 12.0 Hz, 1H), 5.04-4.98 (m, 1H), 4.70 (d, J = 9.2 Hz, 1H), 4.22 (d, J = 14.8, 1H), 2.51-2.45 (m, 1H), 2.32 (s, 4H), 2.22-2.16 (m, 1H), 1.70-1.60 (m, 1H); ¹³C NMR (100 MHz, CDCl₃): δ 157.0, 156.7, 155.9, 137.8, 137.7, 136.2, 132.9, 132.1, 129.2, 128.7, 128.6, 128.2, 128.0, 67.0, 54.0, 49.5, 30.9, 27.1, 21.1; **HRMS** (**ESI**⁺) Calcd. for C₂₁H₂₁ClF₃N₂O₄+H, 433.1739; Found, 433.1741; **IR** (thin film, cm⁻¹) 3439, 3353, 3034, 2950, 2253, 1692,1513, 1209, 1175; **TLC** (70:30 petroleum ether/acetone): R_f = 0.70.



Benzyl (2-(*N*-(4-chlorobenzyl)-2,2,2-trifluoroacetamido) cyclopent-2-en-1-yl)carbamate (15): The title compound was prepared according to General Procedure B using nitrone S6 (23 mg, 0.10 mmol, 1.0 equiv), imidoyl chloride 1 (27 mg, 0.10 mmol, 1.0 equiv) and triethylamine (28 μL, 0.20 mmol, 2.0 equiv) in CH₂Cl₂ (1

mL). Purification by flash chromatography (90:10 hexanes/ethyl acetate) furnished **15** as a colorless oil (31.8 mg, 0.070 mmol, 70%). Analytical data: ¹H NMR (400 MHz, CDCl₃): δ 7.38-7.34 (m, 5H), 7.28 (d, *J* = 8.0 Hz, 2H), 7.19 (d, *J* = 8.0 Hz, 2H), 5.56 (s, 1H), 5.15 (d, *J* = 12.0 Hz, 1H), 5.13 (d, *J* = 14.4 Hz, 1H), 5.07 (d, *J* = 12.0 Hz, 1H), 5.02-4.96 (m, 1H), 4.69 (d, *J* = 9.2 Hz, 1H), 4.25 (d, *J* = 14.8 Hz, 1H), 2.52-2.45 (m, 1H), 2.38-2.32 (m, 1H), 2.24-2.17 (m, 1H), 1.71-1.59 (m, 1H); ¹³C NMR (100 MHz, CDCl₃): δ 157.2, 156.8, 155.9, 137.8, 136.1, 133.9, 133.7, 133.0, 130.1, 128.8, 128.6, 128.3, 128.1, 67.1, 54.0, 49.0, 30.8, 27.2; HRMS (ESI⁺) Calcd. for C₂₂H₂₀ClF₃N₂O₃+H, 453.1194; Found, 453.1191; IR (thin film, cm⁻¹) 3431, 2096, 1690, 1641, 1512, 1265, 1207, 1157, 1051; TLC (70:30 hexanes/ethyl acetate): R_f = 0.48.

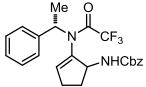


Benzyl (2-(2,2,2-trifluoro-*N*-(4-methoxybenzyl)acetamido)cyclohex-2-en-1-yl)carbamate (16): The title compound was prepared according to General Procedure B using nitrone S7 (23 mg, 0.10 mmol, 1.0 equiv), imidoyl chloride 1 (27 mg, 0.10 mmol, 1.0 equiv) and triethylamine (28 μ L, 0.20 mmol, 2.0 equiv) in CH₂Cl₂ (1 mL). Purification by flash chromatography (90:10 petroleum ether/acetone) furnished 16 as a pale

yellow oil (32 mg, 0.069 mmol, 69%). Analytical data: ¹H NMR (600 MHz, CDCl₃): δ 7.40-7.33 (m, 8H), 7.20 (d, J = 8.4 Hz, 2H), 6.83 (d, J = 8.4 Hz, 2H), 5.41 (s, 1H), 5.29 (d, J = 10.8 Hz, 1H), 5.25 (d, J = 14.4 Hz, 1H), 5.16 (d, J = 12.6 Hz, 1H), 5.06 (d, J = 12.0 Hz, 1H), 4.75 (d, J = 9.6 Hz, 1H), 4.62-4.59 (m, 1H), 4.04 (d, J = 14.4 Hz, 1H), 3.79 (s, 3H), 2.04 (t, J = 16.8 Hz, 2H), 1.94 (d, J = 18.0 Hz, 1H), 1.62-1.59 (m, 3H); ¹³C NMR (150 MHz, CDCl₃): δ 159.2, 156.1, 136.1, 133.4, 132.9, 130.6, 128.6, 128.5, 128.3, 128.1, 127.3, 113.8, 67.1, 55.2, 49.0, 46.1, 30.6, 24.7, 19.9; LRMS (ESI⁺) Calcd. for C₂₄H₂₅F₃N₂O₄+H, 463.18; Found, 463.22; IR (thin film, cm⁻¹) 3356, 3064, 2948, 1691, 1512, 1454, 1323, 1208, 1177; TLC (80:20 petroleum ether/acetone): R_f = 0.30.

Benzyl (5-(tert-butyl)-2-(2,2,2-trifluoro-*N*-(4-methoxybenzyl)acetamido) cyclohex-2-en-1-yl) carbamate (17): The title compound was prepared according to General Procedure B using nitrone S8 (30.5 mg, 0.10 mmol, 1.0 equiv), imidoyl chloride 1 (26.5 mg, 0.01 mmol, 1.0 equiv) and triethylamine (27.9 μ L, 0.20 mmol, 2.0 equiv) in in CH₂Cl₂ (1 mL). Purification by flash chromatography (90:10 hexanes/ethyl acetate) furnished 17 as an off-white solid (21.7 mg, 0.041 mmol, 51%) as a 1.5:1 mixture of separable

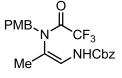
diastereomers. Analytical data: ¹H NMR (600 MHz, CDCl₃): (Major Diastereomer) δ 7.37-7.34 (m, 5H), 7.17 (d, *J* = 6.6 Hz, 2H), 7.10 (d, *J* = 8.4 Hz, 1H), 6.83 (d, *J* = 7.2 Hz, 3H), 5.52 (br s, 1H), 5.14 (d, *J* = 12.0 Hz, 1H), 5.08 (d, *J* = 13.2 Hz, 1H), 5.00-4.66 (m, 1H), 4.66 (br s, 1H), 4.15 (d, *J* = 15.0 Hz, 1H), 3.79 (s, 3H), 3.12-3.10 (m, 1H), 2.10-2.07 (m, 1H), 1.97-1.95 (m, 1H), 1.72-1.68 (m, 1H), 1.40-1.30 (m, 1H), 1.20-1.15 (m, 1H), 0.85 (s, 9H); (Minor Diastereomer) δ 7.39-7.35 (m, 5H), 7.21 (d, *J* = 8.4 Hz, 2H), 6.84 (d, *J* = 8.4 Hz, 2H), 5.38-5.36 (m, 1H), 5.22 (d, *J* = 14.4 Hz, 1H), 5.17 (d, *J* = 12.0 Hz, 1H), 5.06 (d, *J* = 12.0 Hz, 1H), 4.64 (s, 2H), 3.99 (d, *J* = 14.4 Hz, 1H), 3.80 (s, 3H), 2.20 (d, *J* = 12.0 Hz, 1H), 1.99-1.96 (m, 1H), 1.83-1.78 (m, 1H), 1.42-1.41 (m, 1H), 1.21-1.17 (m, 1H), 0.86 (s, 9H); ¹³C NMR (150 MHz, CDCl₃): (Minor Diastereomer) δ 159.2, 156.2, 136.1, 133.7, 132.2, 130.6, 128.6, 128.3, 128.1, 127.4, 113.8, 67.2, 55.2, 49.1, 48.1, 43.2, 32.6, 32.1, 26.9, 26.3; HRMS (ESI⁺) Calcd. for C₂₈H₃₃F₃N₂O₄+Na, 541.2291; Found, 541.2290; IR (thin film, cm⁻¹) 3487, 3035, 2962, 1803, 1712, 1612, 1498, 1455, 1251; TLC (70:30 hexanes/ethyl acetate): R_f = 0.49.



Benzyl (2-(2,2,2-trifluoro-*N*-((*S*)-1-phenylethyl)acetamido)cyclopent -2-en-1-yl)carbamate (18): The title compound was prepared according to General Procedure B using nitrone **S9** (40 mg, 0.20 mmol, 1.0 equiv), imidoyl chloride 1 (53 mg, 0.20 mmol, 1.0 equiv) and triethylamine (56 μL, 0.40 mmol, 2.0 equiv) in CH₂Cl₂ (2 mL).

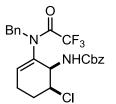
Purification by flash chromatography (90:10 hexanes/ethyl acetate) furnished **18** as a pale yellow oil which was a 1:1 mixture of separable diastereomers (Diastereomer 1: 20.2 mg, 0.047 mmol,

23%; Diastereomer 2: 19 mg, 0.043, 22%). Analytical data: ¹H NMR (400 MHz, CDCl₃): (Diastereomer 1) δ 7.38-7.31 (m, 10H), 5.73-5.68 (m, 1H), 5.48 (br s, 1H), 5.13 (d, *J* = 12.4 Hz, 1H), 5.09 (d, *J* = 8.4 Hz, 1H), 4.84 (d, *J* = 7.2 Hz, 1H), 2.35-2.21 (m, 2H), 1.99 (br s, 1H), 1.63 (d, *J* = 7.2 Hz, 3H), 1.59-1.48 (m, 1H); (Diastereomer 2) δ 7.39-7.27 (m, 10H), 5.84 (s, 1H), 5.45-5.40 (m, 1H), 5.01 (s, 2H), 4.60 (br s, 1H), 4.30 (br s, 1H), 2.38-2.30 (m, 4H), 1.62 (d, *J* = 6.8 Hz, 3H); ¹³C NMR (150 MHz, CDCl₃): (Diastereomer 1) 156.6, 156.4, 155.9, 138.1, 136.8, 136.2, 135.9, 128.6, 128.4, 128.3, 128.2, 67.0, 56.5, 32.1, 26.8, 16.3; (Diastereomer 2) δ 155.3, 138.3, 137.5, 136.2, 134.8, 128.5, 128.2, 128.1, 127.6, 66.8, 56.4, 55.2, 32.0, 27.1, 16.4; HRMS (ESI⁺) Calcd. for C₂₃H₂₃F₃N₂O₃+H, 433.1740; Found, 433.1750; IR (thin film, cm⁻¹) 3422, 3054, 2986, 2305, 1685, 1508, 1420, 1265, 1206, 1155; TLC (70:30 hexane/ethyl acetate): R_f = Diastereomer 1: 0.55; Diastereomer 2: 0.45.



(Z)-Benzyl (2-(2,2,2-trifluoro-*N*-(4-methoxybenzyl)acetamido) prop-1en-1-yl)carbamate (19): The title compound was prepared according to General Procedure B using nitrone S10 (20 mg, 0.10 mmol, 1.0 equiv), imidoyl chloride 1 (27 mg, 0.10 mmol, 1.0 equiv) and triethylamine (28 μ L, 0.20 mmol, 2.0 equiv) in CH₂Cl₂ (1 mL). Purification by flash

chromatography (95:5 to 90:10 petroleum ether/acetone) furnished **19** as a pale yellow oil (19 mg, 0.045 mmol, 45%). Analytical data: ¹**H NMR** (600 MHz, CDCl₃): δ 7.37-7.29 (m, 3H), 7.24 (d, *J* = 7.2 Hz, 2H), 6.83 (d, *J* = 8.4 Hz, 2H), 6.52 (d, *J* = 10.8 Hz, 1H), 5.62 (d, *J* = 11.4 Hz, 1H), 5.12 (d, *J* = 13.8 Hz, 1H), 5.03 (d, *J* = 12.0 Hz, 1H), 4.96 (d, *J* = 12.0, 1H), 4.02 (d, *J* = 13.8 Hz, 1H), 3.68 (s, 3H), 1.85 (s, 3H); ¹³C NMR (150 MHz, CDCl₃): δ 159.9, 152.7, 135.4, 130.7, 128.6, 128.3, 127.9, 127.8, 123.7, 114.4, 113.7, 67.4, 55.2, 49.8, 18.9; **HRMS (ESI**⁺) Calcd. for C₂₁H₂₁ClF₃N₂O₄+Na,445.1351; Found, 445.1354; **IR** (thin film, cm⁻¹) 3415, 2958, 2253, 1693, 1513, 1213, 1173, 1034; **TLC** (70:30 petroleum ether/acetone): R_f = 0.70.

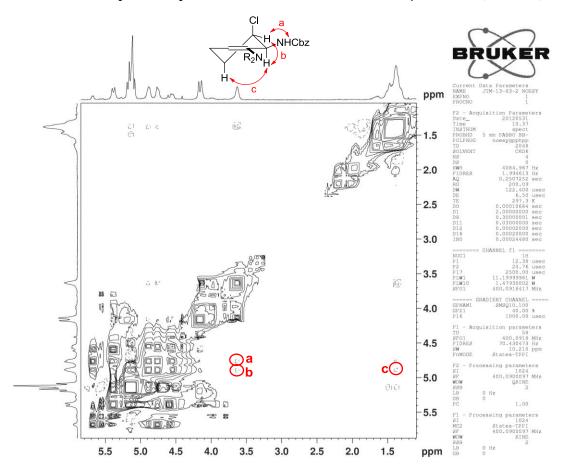


Benzyl (2-(*N*-benzyl-2,2,2-trifluoroacetamido)-1-chlorocyclohex-2-en-1yl)carbamate (20): The title compound was prepared according to General Procedure B using nitrone $S11^6$ (23 mg, 0.10 mmol, 1.0 equiv), imidoyl chloride 1 (27 mg, 0.10 mmol, 1.0 equiv) and triethylamine (28 µL, 0.20 mmol, 2.0 equiv) in CH₂Cl₂ (1 mL). Purification by flash chromatography (90:10 petroleum ether/acetone) furnished 20 as a pale yellow oil (39 mg,

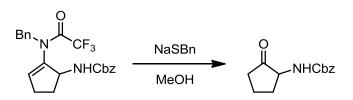
0.084 mmol, 84%). Analytical data: ¹**H NMR** (600 MHz, CDCl₃): δ 7.39-7.30 (m, 11H), 5.53 (s, 1H), 5.27 (d, *J* = 15.0 Hz, 1H), 5.19 (d, *J* = 12.0 Hz, 1H), 5.11 (d, *J* = 6.0 Hz, 1H), 5.09 (d, *J* = 7.8 Hz, 1H), 4.90 (d, *J* = 9.0 Hz, 1H), 4.74-4.67 (m, 1H), 4.11 (d, *J* = 14.4 Hz, 2H), 2.23 (br s, 1H), 2.16-2.13 (m, 1H), 2.07-2.05 (m, 1H), 2.01-1.96 (m, 1H); ¹³**C NMR** (150 MHz, CDCl₃): δ 156.1, 135.8, 134.8, 132.8, 130.5, 129.3, 128.8, 128.7, 128.6, 128.6, 128.5, 128.4, 128.1, 127.9, 67.5, 58.9, 53.0, 49.7, 27.6, 22.0; **HRMS** (**ESI**⁺) Calcd. for C₂₃H₂₂ClF₃N₂O₃+Na, 467.1271; Found, 467.1361; **IR** (thin film, cm⁻¹) 3439, 3068, 2960, 2253, 1692, 1500, 1442, 1213, 1158; **TLC** (70:30 petroleum ether/acetone): R_f = 0.50.

¹H NOESY analysis of 20:

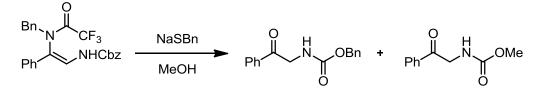
The *cis* chloro/amino stereochemistry was supported by NOESY spectroscopy. The β -methine (R₂CHCl) showed two nOe's: a strong interaction (a) with the amino methine (RNHCbz), and interaction (b) with the allylic methine proton (R₂CHNHCbz). Additionally, the allylic methine displayed an interaction (c) with the γ -alkyl methylene, an interaction that would not be observed if the allylic methine occupied the pseudoequitorial position – the only orientation that would support a *trans* relationship based upon the dual nOe observed with the β -methine (R₂CHCl).



Secondary Transformations:

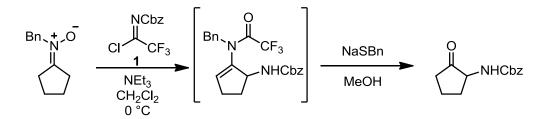


Benzyl (2-oxocyclopentyl)carbamate (21): A oven-dried 4-mL vial equipped with magnetic stir bar was charged with α'-amino enamide **3** (0.065 g, 0.15 mmol, 1.0 equiv). Methanol (1.0 mL) was added, followed by freshly prepared sodium benzylthiolate (0.112 g, 0.77 mmol, 5.0 equiv). The vial was sealed with a Teflon-lined cap and the reaction was stirred at rt for 14 h. Volatiles were removed *in vacuo* and the crude product was purified via flash chromatography (90:10 to 80:20 petroleum ether/acetone) to provide **21** as a pale yellow oil (29 mg, 85%). Analytical data: ¹H NMR (600 MHz, CDCl₃): δ 7.38-7.31 (m, 5H), 5.22 (br s, 1H), 5.11 (s, 2H), 3.99-3.96 (m, 1H), 2.64-2.62 (m, 1H), 2.42 (dd, *J* = 19.8, 9.0 Hz, 1H), 2.21-2.14 (m, 1H), 2.09-2.04 (m, 1H), 1.86-1.84 (m, 1H), 1.67-1.63 (m, 1H); ¹³C NMR (150 MHz, CDCl₃): δ 214.6, 156.2, 136.2, 128.5, 128.2, 128.1, 67.0, 59.2, 34.8, 30.2, 17.8; MS (ESI⁺) Calcd. for C₁₃H₁₅NO₃+Na, 256.10; Found, 256.10; IR (thin film, cm⁻¹) 3414, 3055, 2978, 2885, 2305, 1719, 1509, 1455, 1266, 1063; TLC (60:40 petroleum ether/acetone): R_f = 0.40.

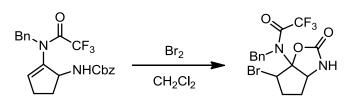


Benzyl (2-oxo-2-phenylethyl)carbamate and Methyl (2-oxo-2-phenylethyl)carbamate (22): A oven-dried 20-mL scintillation vial equipped with magnetic stir bar was charged with enediamine 7 (0.160 g, 0.346 mmol, 1.0 equiv). Methanol (4.0 mL) was added, followed by freshly prepared sodium benzylthiolate (0.202 g, 1.38 mmol, 4.0 equiv). The vial was sealed with a Teflon-lined cap and the reaction was stirred at rt for 2 h. Volatiles were removed in vacuo. THF (1.5 mL) and 1 M HCl (aq.) were added to the vial and the reaction was allowed to stir at rt for 0.5 h. The organic layer was extracted with Et₂O (3 x 2 mL) and dried with MgSO₄. Volatiles were removed in vacuo and the crude product was purified via flash chromatography (85:15 to 75:25 hexanes/ethyl acetate) to provide two products. Cbz: colorless oil, 38 mg, 41%, OMe: white solid, 7 mg, 11%. Analytical data: Cbz Product: ¹H NMR (400 MHz, CDCl₃): δ 7.96 (d, J = 7.6 Hz, 2H), 7.62 (t, J = 7.6 Hz, 1H), 7.49 (t, J = 7.6 Hz, 2H), 5.85 (br s, 1H), 5.16 (s, 2H), 4.73-4.69 (m, 2H); ¹³C NMR (100 MHz, CDCl₃): δ 193.2, 156.2, 136.3, 134.0, 128.9, 128.5, 128.1, 128.0, 127.8, 126.9, 67.0, 47.8; MS (ESI⁺) Calcd. for C₁₆H₁₅NO₃+H, 270.10; Found, 270.07; IR (thin film, cm⁻¹) 3417, 3061, 1692, 1511, 1354, 1266, 1218, 1057; TLC (70:30 hexanes/ethyl acetate): $R_f = 0.30$. OMe Product: mp 77-82 °C; ¹H NMR (400 MHz, CDCl₃): δ 7.96 (d, J = 7.6 Hz, 2H), 7.62 (t, J = 7.6 Hz, 1H), 7.49 (t, J = 7.6 Hz, 2H), 5.74 (br s, 1H), 4.71 (d, J = 4.4 Hz, 2H), 3.72 (s, 3H); ¹³C NMR (100 MHz, CDCl₃): δ 194.1, 156.9, 134.3, 134.0,

128.9, 127.8, 52.4, 47.8; **MS** (**ESI**⁺) Calcd. for $C_{10}H_{11}NO_3+Na$, 216.06; Found, 216.01; **IR** (thin film, cm⁻¹) 3426, 1692, 1518, 1536, 1265, 737, 701; **TLC** (70:30 hexanes/ethyl acetate): $R_f = 0.20$.



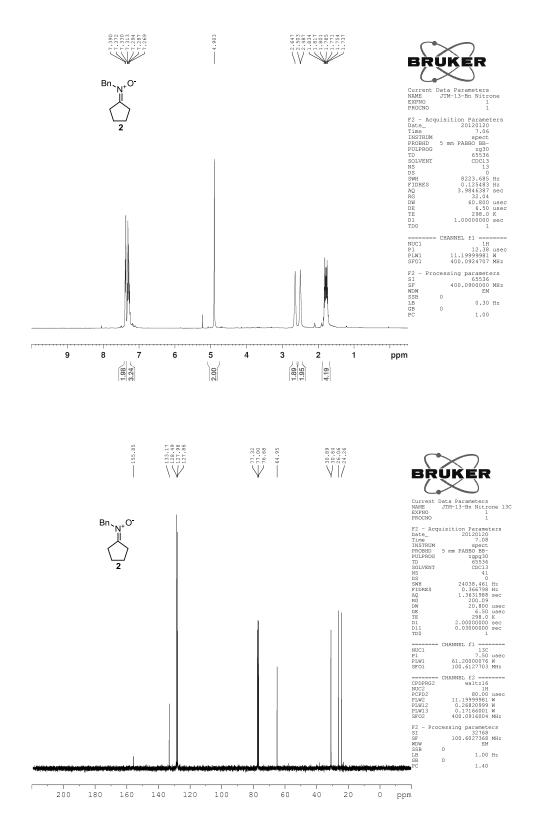
Benzyl (2-oxocyclopentyl)carbamate (21): The title compound was prepared according to General Procedure B using nitrone **2** (71 mg, 0.377 mmol, 1.0 equiv), imidoyl chloride **1** (100 mg, 0.377 mmol, 1.0 equiv) and triethylamine (100 μ L, 0.755 mmol, 2.0 equiv) in CH₂Cl₂ (2 mL). Upon completion, volatiles were removed *in vacuo*. Methanol was immediately added to the vial (3.0 mL) followed by the addition of freshly prepared sodium benzylthiolate (0.165 g, 1.13 mmol, 3.0 equiv). The vial was sealed with a Teflon-lined cap and the reaction was stirred at rt for 14 h. Volatiles were removed *in vacuo* and the crude product was purified via flash chromatography (90:10 to 80:20 petroleum ether/acetone) to provide the desired product as a pale yellow oil (58 mg, 67%).

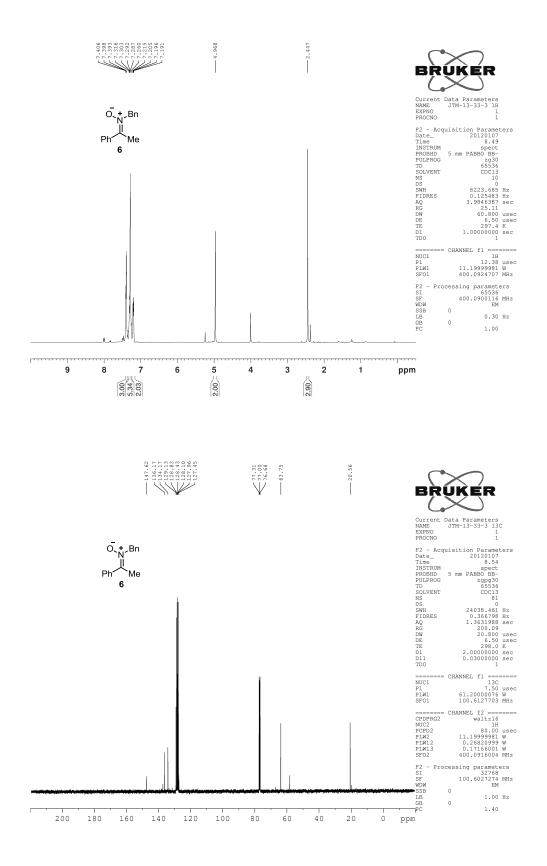


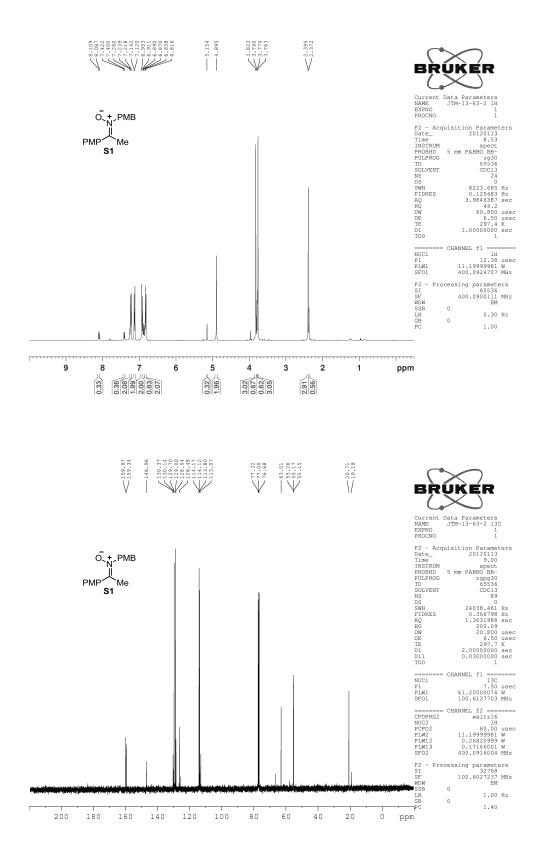
N-benzyl-N-(6-bromo-2-oxohexahydro-2H-cyclopenta[d]oxazol-6a-yl)-2,2,2-

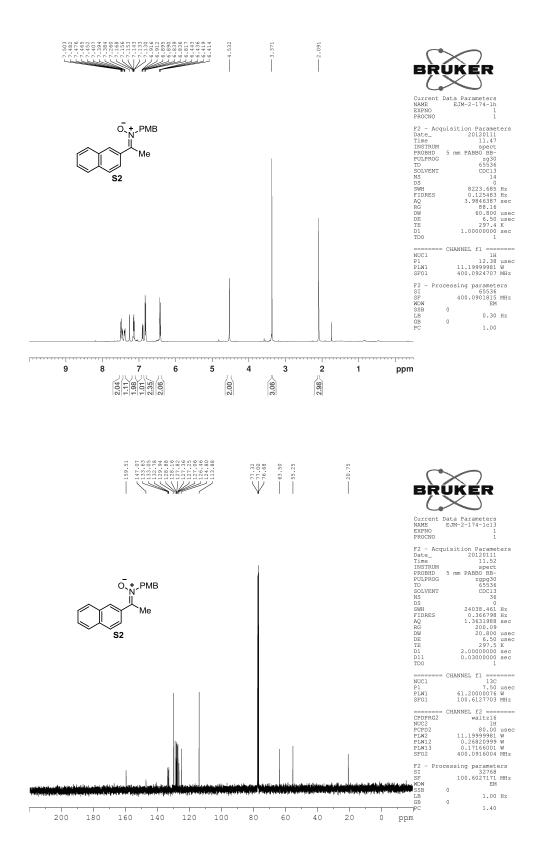
trifluoroacetamide (23): A oven-dried 4-mL scintillation vial equipped with magnetic stir bar was charged with α '-amino enamide **3** (0.020 g, 0.048 mmol, 1.0 equiv). The vial was purged with nitrogen and dichloromethane (1.2 mL) was added. Bromine (0.007 mL, 0.14 mmol, 3.0 equiv) was then added dropwise and the reaction was stirred at 23 °C for 45 min. Upon completion, volatiles were removed *in vacuo*. The crude product was purified via flash chromatography (90:10 to 80:20 petroleum ether/acetone) to provide the desired product as a white solid with 2.5:1 diastereomeric ratio (15 mg, 79%). Analytical data: **mp** 176.5-178 °C; ¹**H NMR** (600 MHz, CDCl₃): (Major Diastereomer) δ 7.38-7.22 (m, 5H), 6.01 (s, 1H), 5.49 (s, 1H), 5.01 (d, *J* = 19.8 Hz, 1H), 4.68 (d, *J* = 18.0 Hz, 1H), 4.28 (d, *J* = 4.2 Hz, 1H), 2.38-2.21 (m, 3H), 1.66 (d, *J* = 12.6 Hz, 1H); ¹³**C NMR** (150 MHz, CDCl₃): (Major Diastereomer) δ 156.7, 135.5, 129.1, 128.6, 128.1, 125.5, 116.8, 114.9, 105.2, 57.8, 56.2, 50.6, 32.5, 31.4; **HRMS (ESI**⁺) Calcd. for C₁₅H₁₄BrF₃N₂O₃ +Na, 429.0038; Found, 429.0039; **IR** (thin film, cm⁻¹) 3054, 2986, 2305, 1778, 1719, 1292, 1209, 1149, 740; **TLC** (60:40 hexanes/ethyl acetate): $R_f = 0.30$.

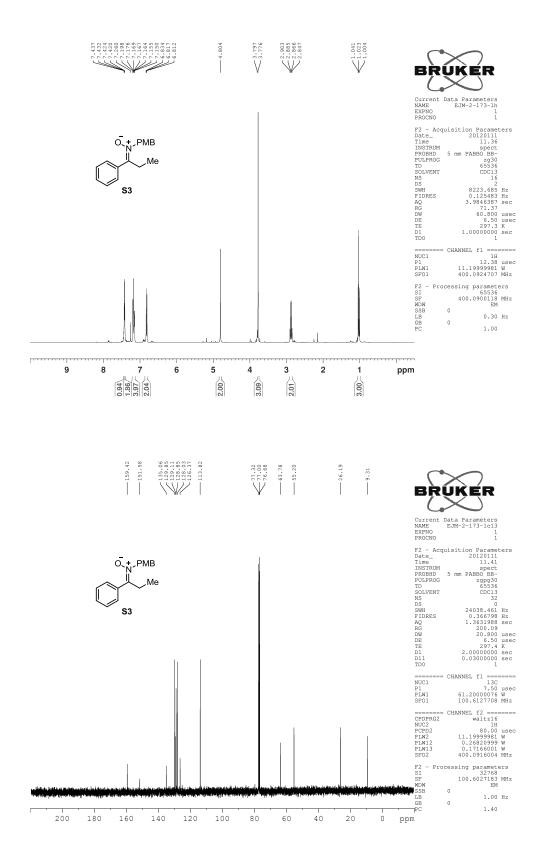
¹H, ¹³C NMR Spectra:

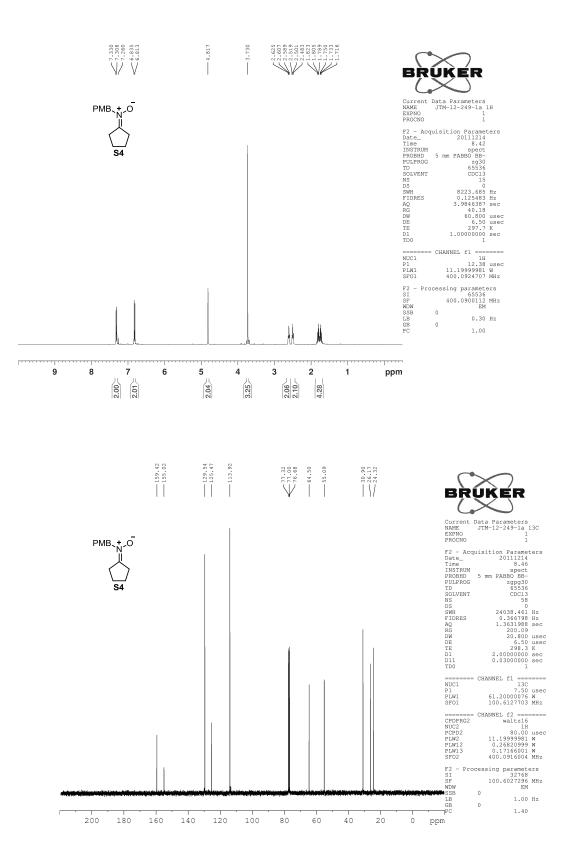


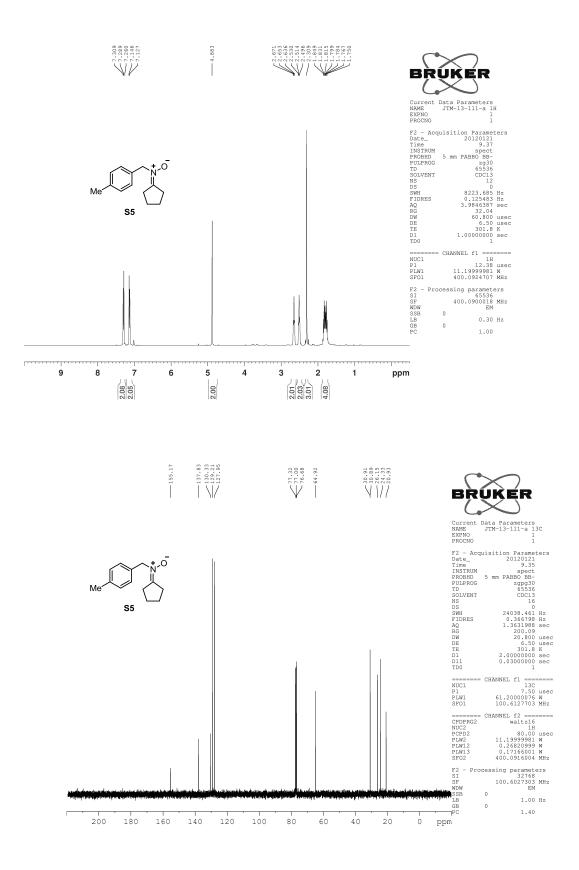


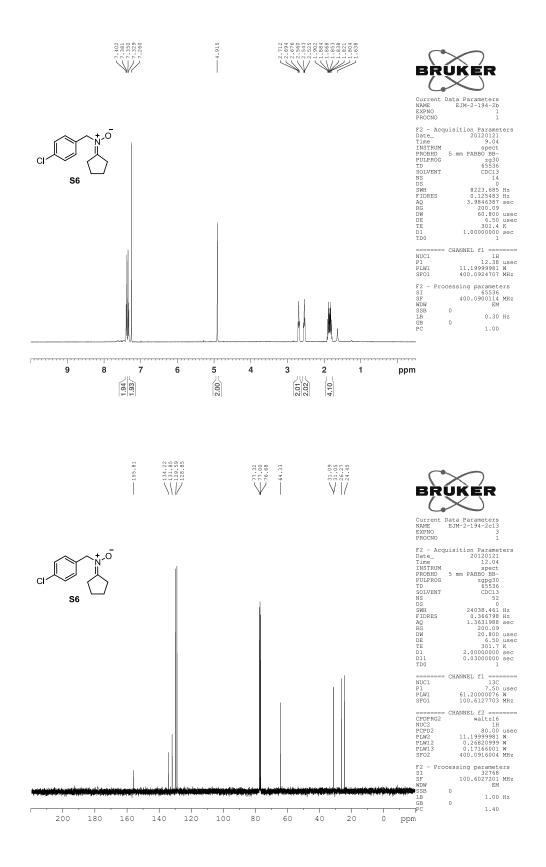


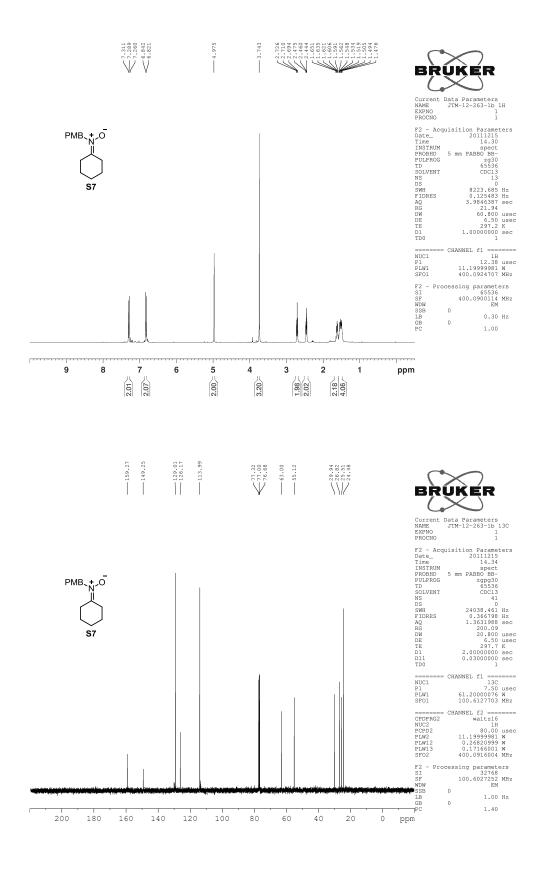


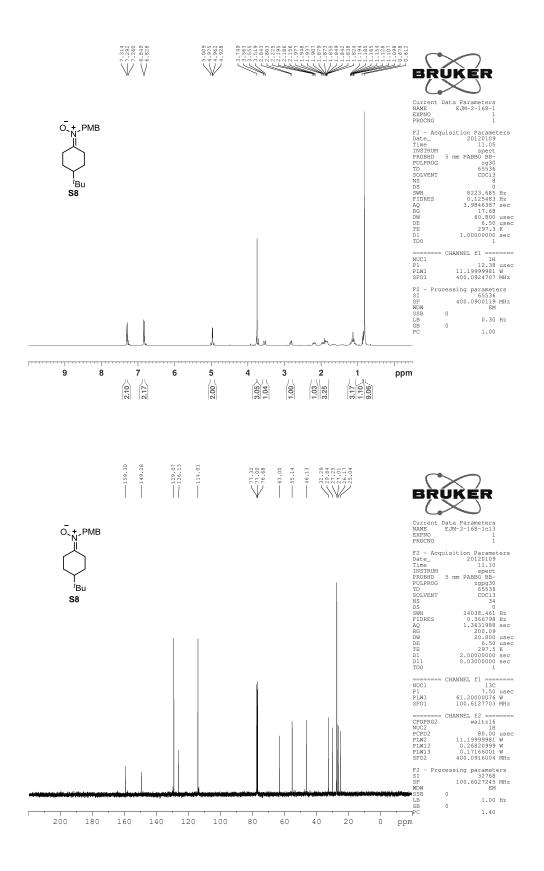


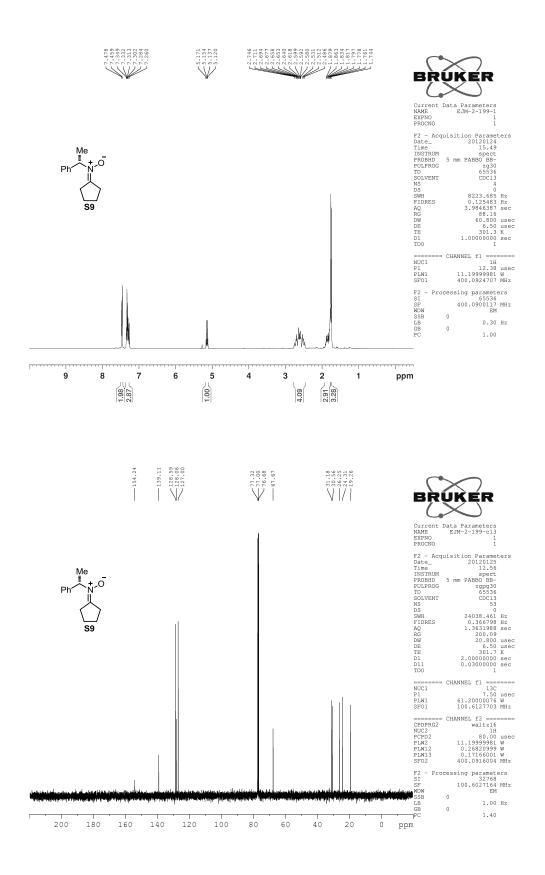


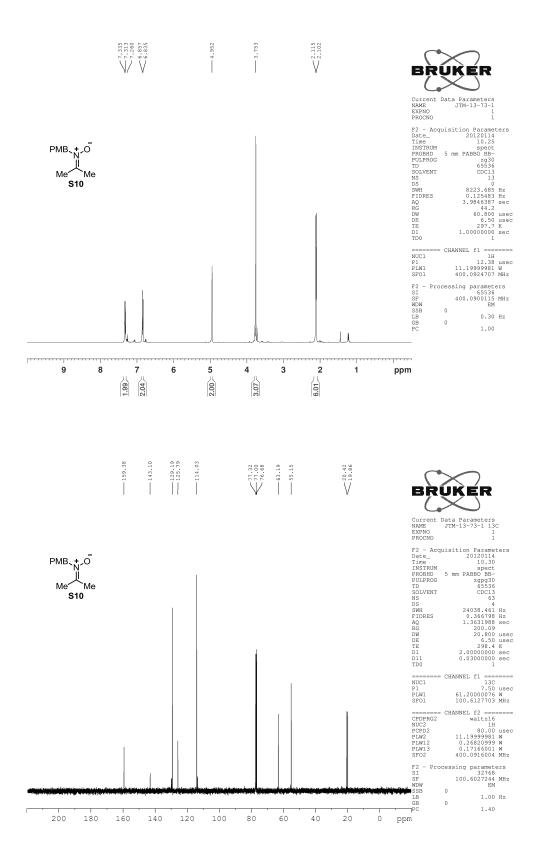




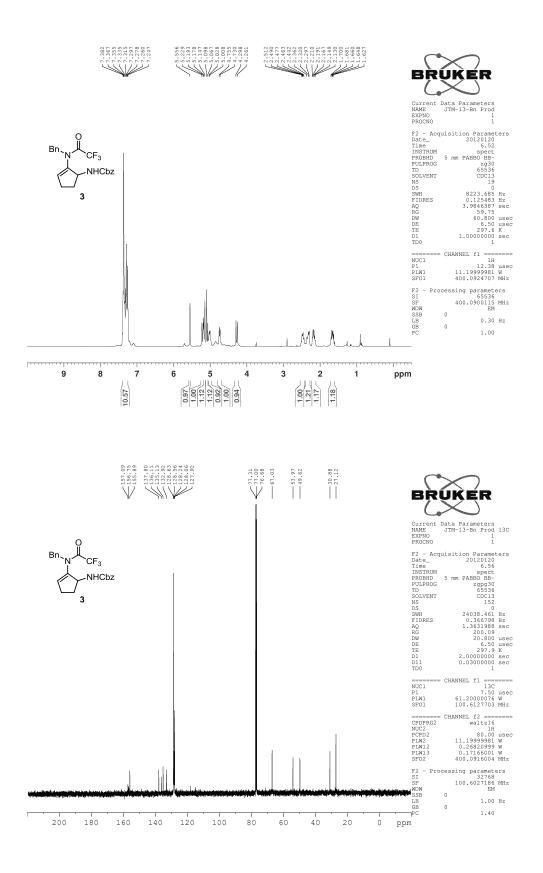




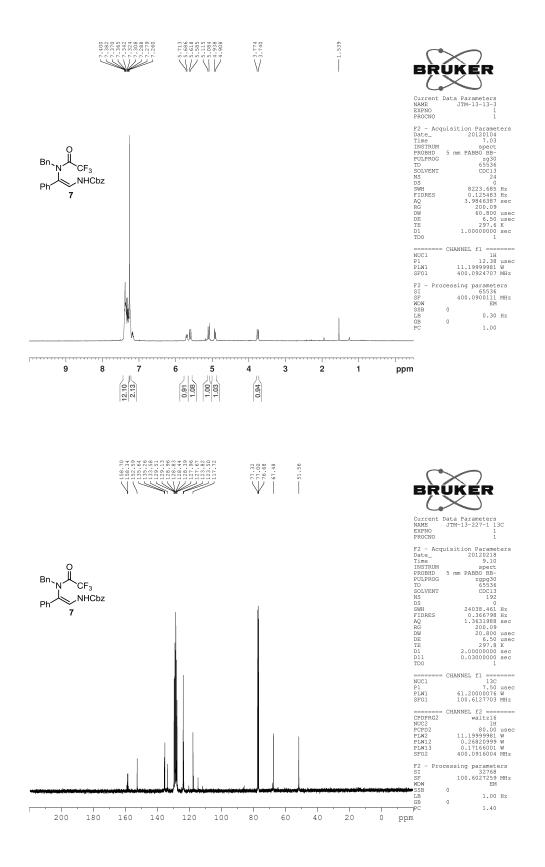


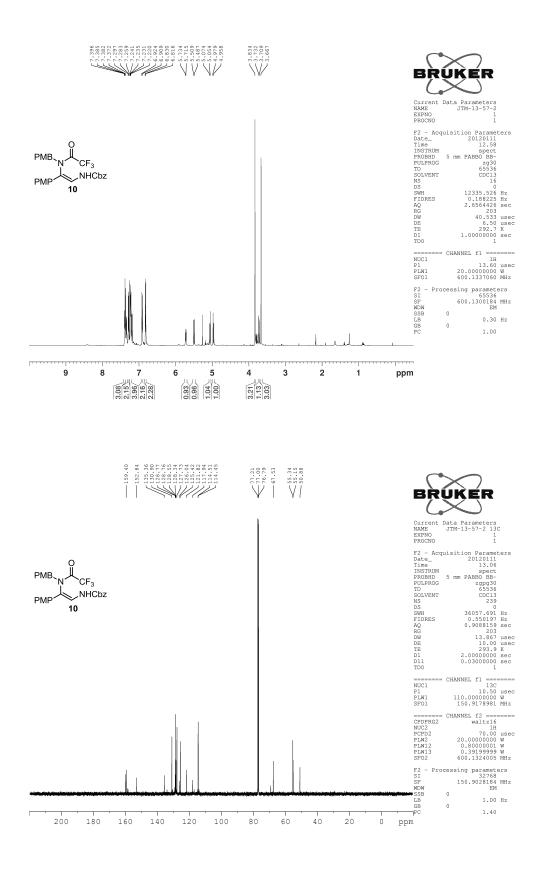


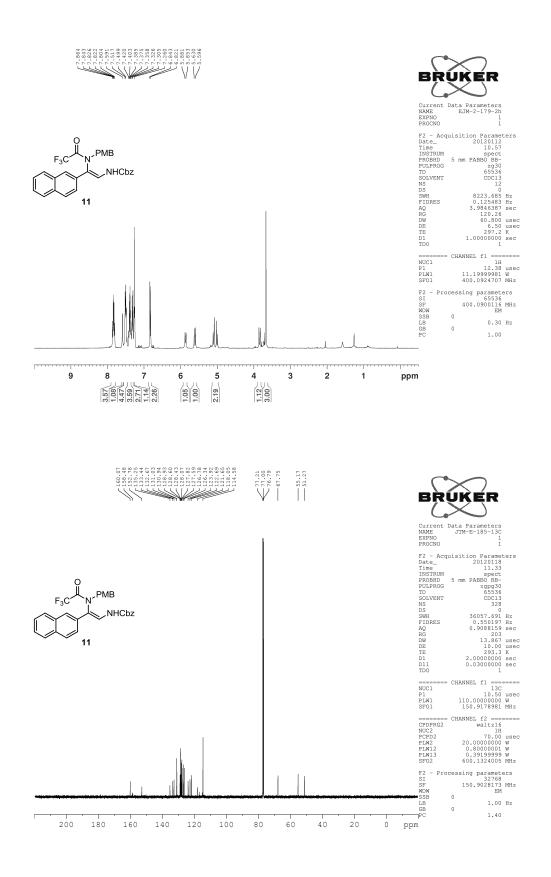
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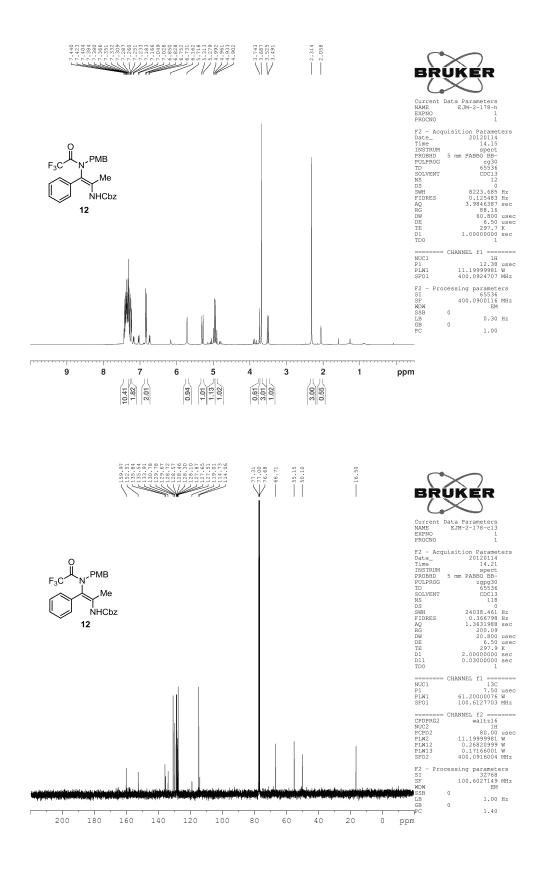


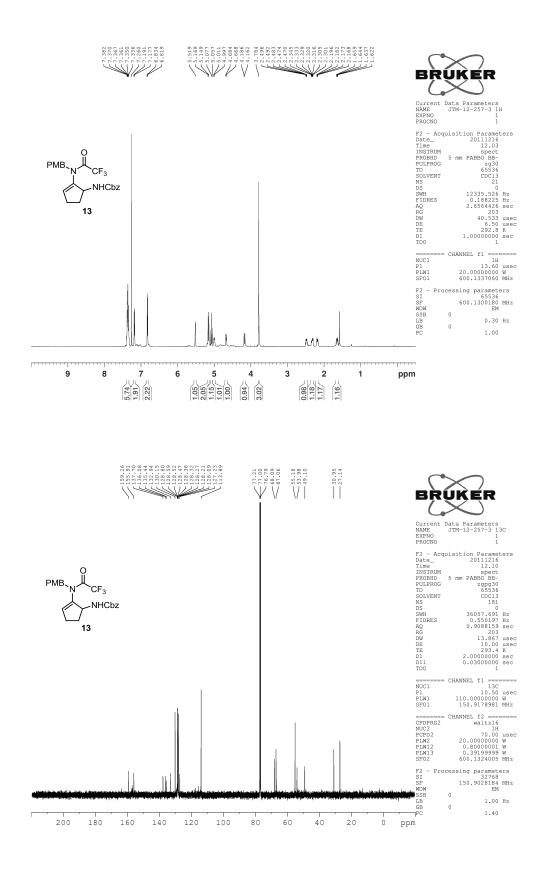
S28

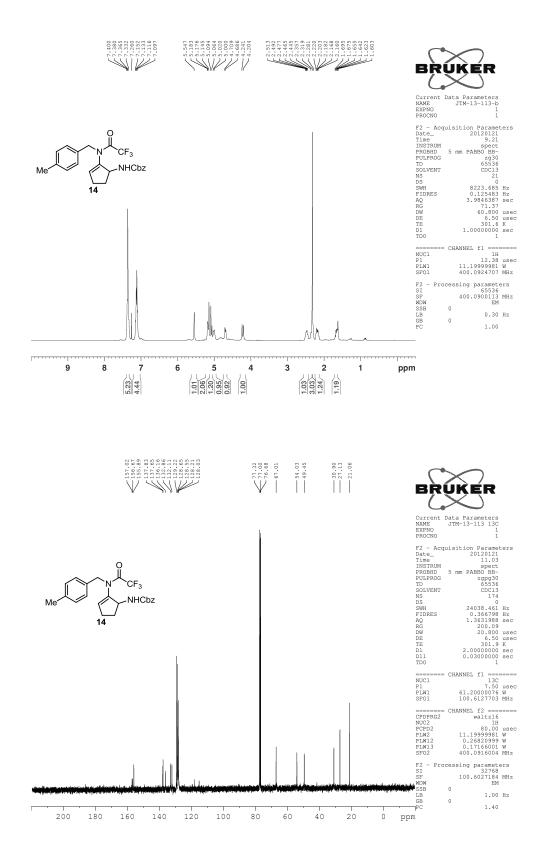


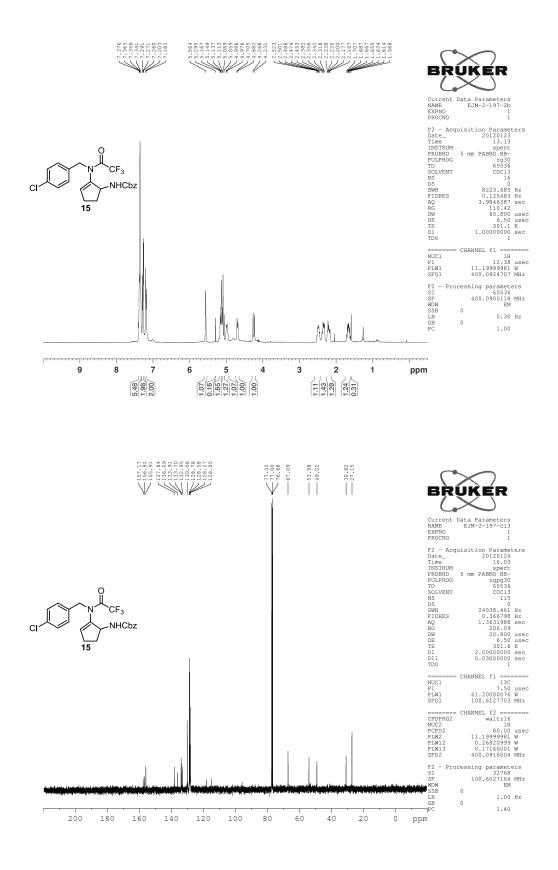


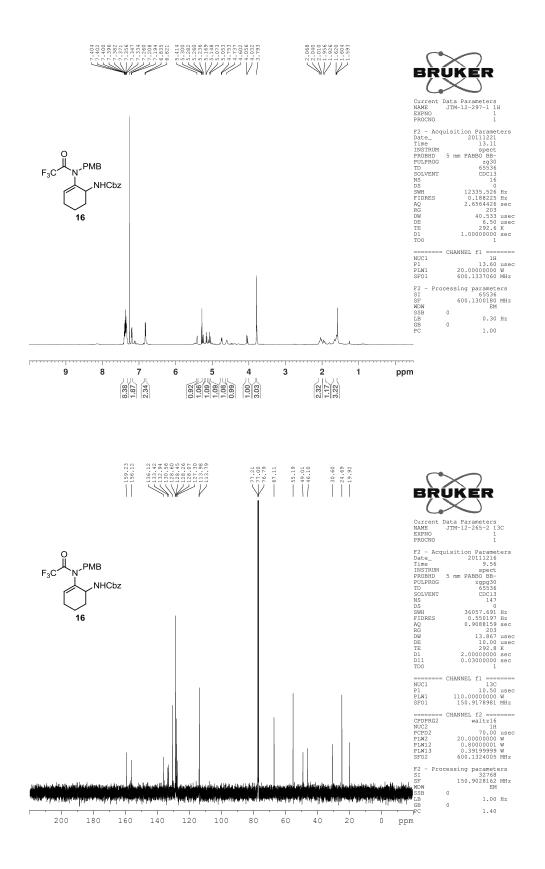


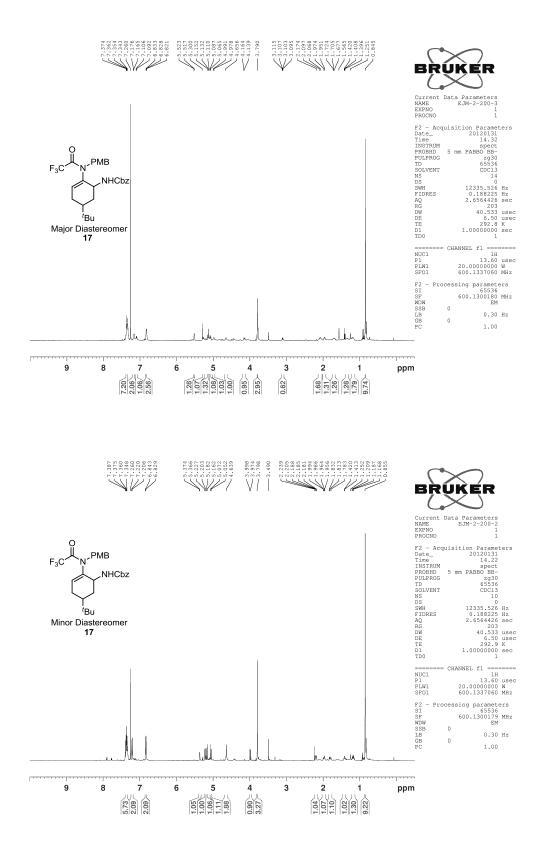


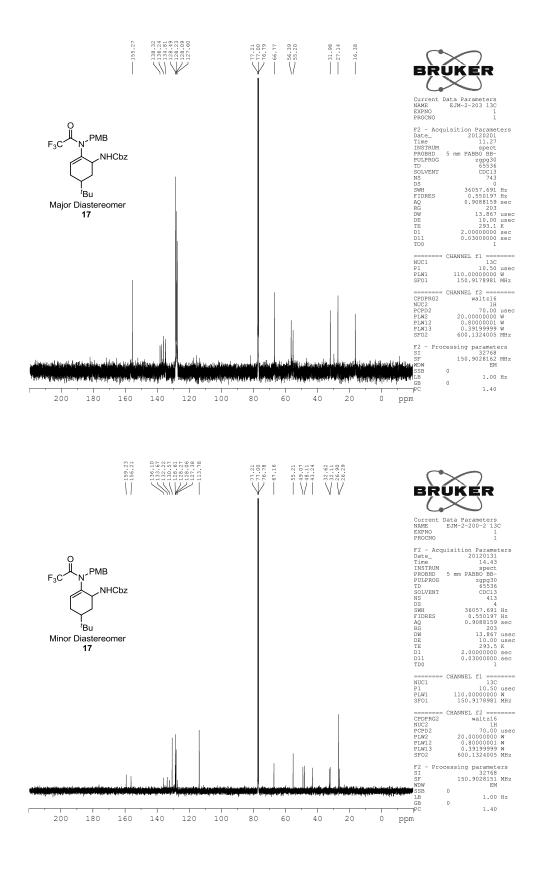


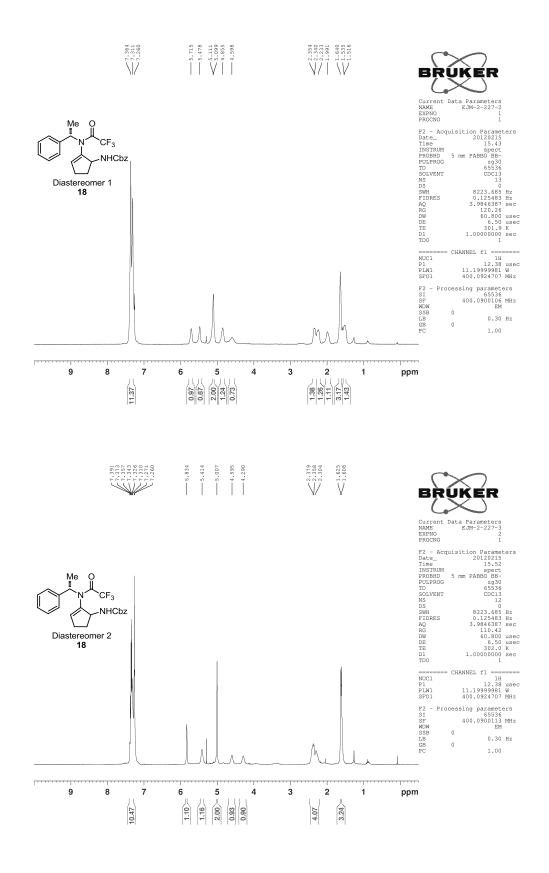


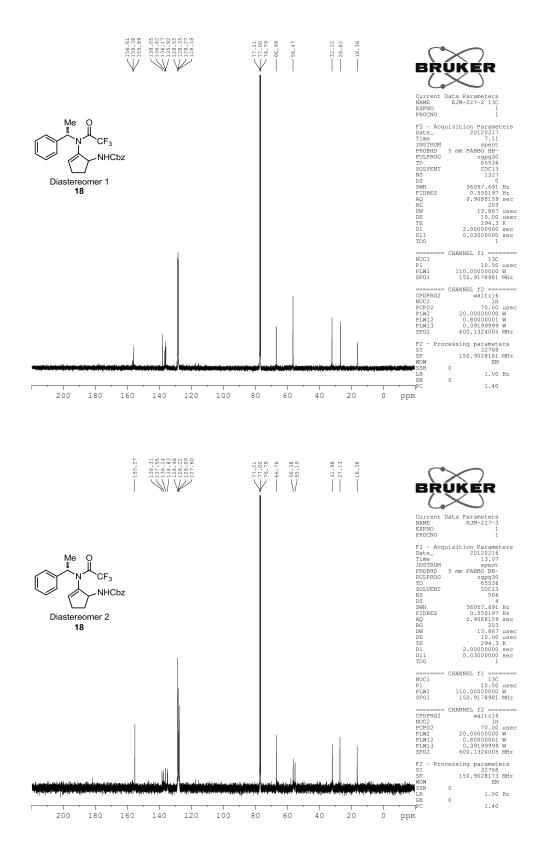


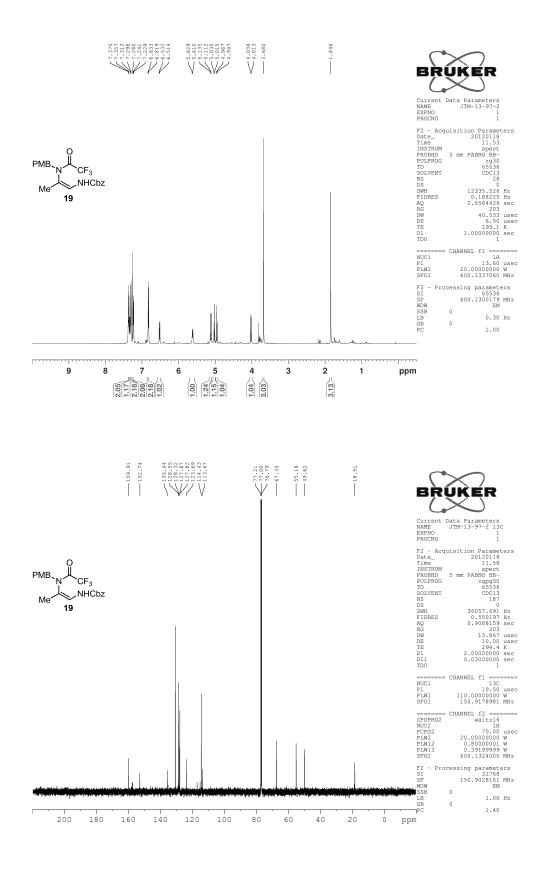


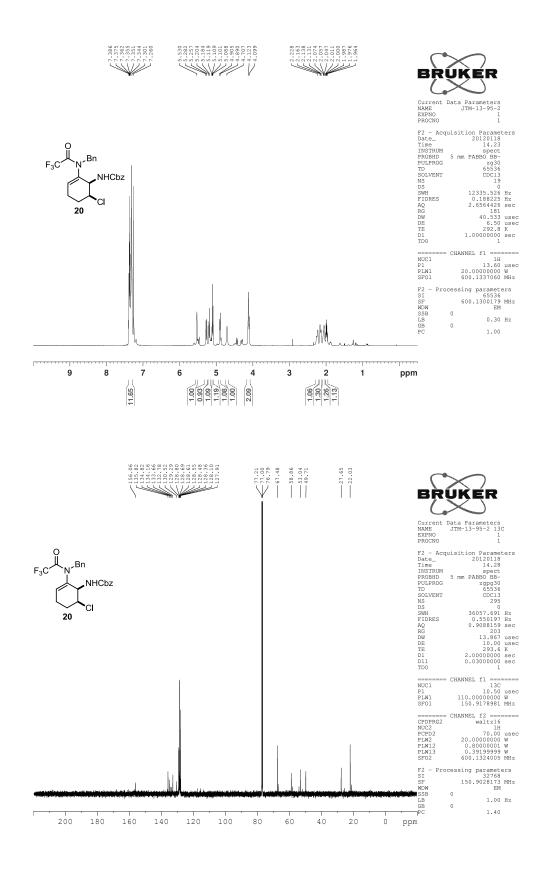


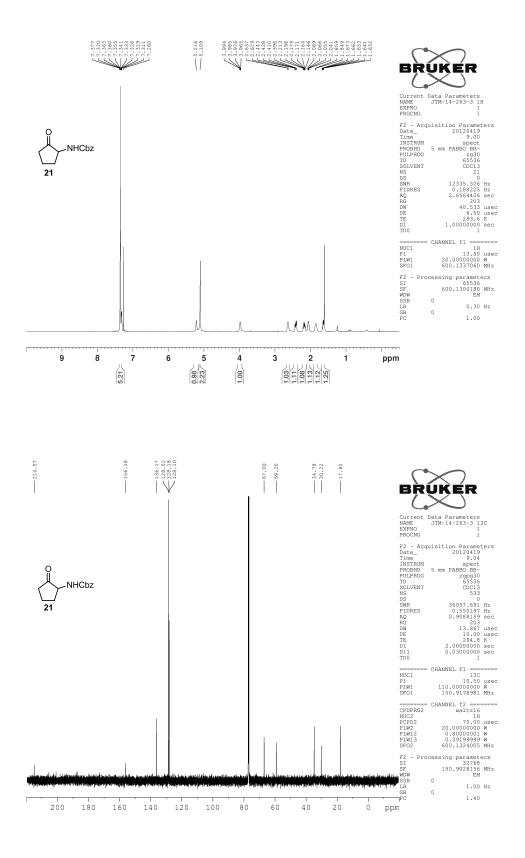


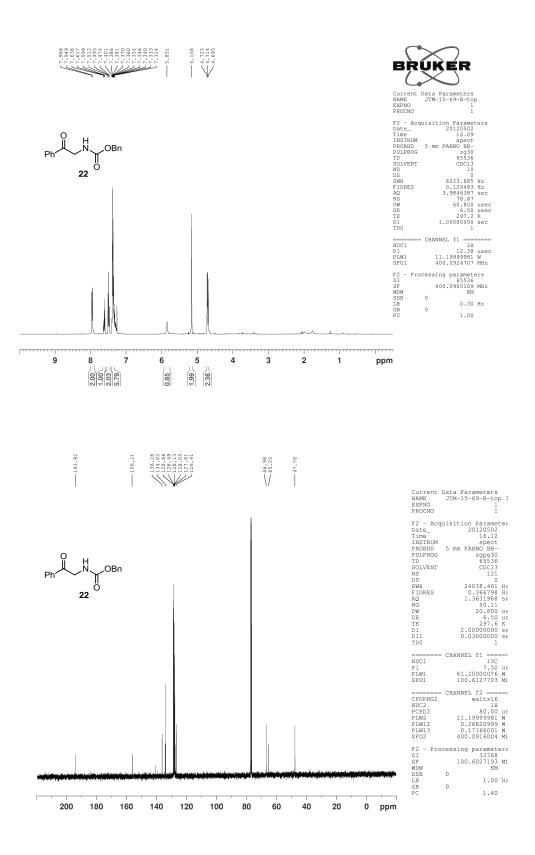












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