

Asymmetric Hydrogenation of TIPS-Protected Oximes with Chiral Boranes

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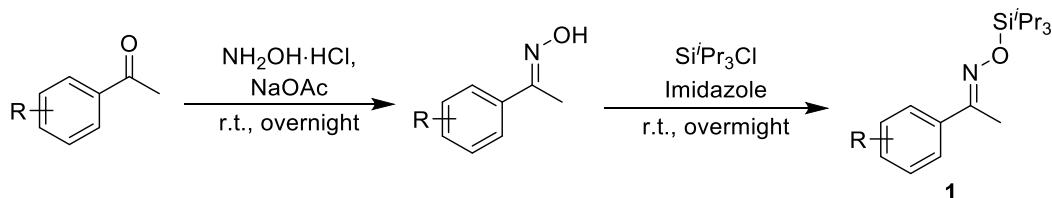
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Supporting Information

General consideration: All air-sensitive compounds were handled under an atmosphere of argon or in a nitrogen-filled glovebox. ^1H NMR, ^{13}C NMR and ^{19}F NMR spectra were recorded on Bruker AV 300, 400 or 500 at ambient temperature with CDCl_3 as solvent and TMS as internal standard. Chemical shifts (δ) were given in ppm, referenced to the residual proton resonance of TMS (0), to the carbon resonance of the CDCl_3 (77.23). Coupling constants (J) were given in Hertz (Hz). IR spectrums were recorded on Perkin-Elmer-983 spectrometer. Column chromatography was performed on silica gel (200-300 mesh). All solvents were purified by conventional methods, distilled before use. Commercially available reagents were used without further purification.

Representative procedure for the synthesis of TIPs-protected oxime 1f



A glass vial was charged with acetophenone (0.60 g, 5.0 mmol), hydroxylamine hydrochloride (0.35 g, 5.0 mmol, 1.0 equiv.), sodium acetate (0.41 g, 5.0 mmol, 1.0 equiv.), and methanol (10 mL) without exclusion of oxygen or moisture. After stirring at room temperature overnight, the reaction mixture was diluted with water. The aqueous phase was extracted with CH_2Cl_2 (3×10 mL) and the combined organic phases were dried over Na_2SO_4 and filtered. Evaporation of the solvent under reduced pressure yielded the crude oximes which were used without further purification.

A flame-dried Schlenk flask was charged with the above obtained oxime (0.61 g, 4.5 mmol), triisopropylsilyl chloride (1.2 mL, 5.4 mmol, 1.2 equiv.), and CH_2Cl_2 (5 mL). Imidazole (0.62 g, 9 mmol, 2.0 equiv.) in CH_2Cl_2 (10 mL) was added dropwise, and the reaction mixture was stirred at room temperature overnight. After evaporation of the solvent, the residue was purified by flash

column chromatography on silica gel using petroleum ether/EA = 20/1 as eluent to give the desired product **1f** (1.19 g, 90% yield).

J. Mohr and M. Oestreich, *Angew. Chem. Int. Ed.*, 2014, **53**, 13278-13281.

Representative procedure for the preparation of racemic products (rac-2f): To a stainless-steel autoclave were added $\text{B}(\text{C}_6\text{F}_5)_3$ (15.3 mg, 0.03 mmol), (*E*)-1-phenylethan-1-one *O*-triisopropylsilyl oxime (**1f**) (87.5 mg, 0.30 mmol), and dry toluene (1.5 mL) in a nitrogen atmosphere glovebox. After being sealed, the autoclave was purged three times with H_2 and the final pressure of hydrogen was adjusted to 4 MPa. The resulting mixture was stirred at 60 °C in an oil bath for 18 h. The solvent was evaporated under reduced pressure and the crude residue was purified by column chromatography (petroleum ether/EA = 20/1) on silica gel to give *N*-(1-phenylethyl)-*O*-(triisopropylsilyl)hydroxylamine (**rac-2f**) as a colorless oil.

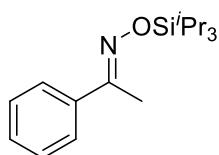
Representative procedure for asymmetric hydrogenation of *O*-triisopropylsilyl oxime **1f (Scheme 3):** To a stainless-steel autoclave, were added $\text{HB}(\text{C}_6\text{F}_5)_2$ (**4**) (10.4 mg, 0.03 mmol), chiral diene **3f** (16.6 mg, 0.015 mmol), and dry toluene (1.5 mL) in a nitrogen atmosphere glovebox. The resulting mixture was stirred for 10 min at room temperature followed by addition of TIPS-protected oxime **1f** (87.5 mg, 0.30 mmol). After being sealed, the autoclave was purged three times with H_2 and the final pressure of hydrogen was adjusted to 40 bar. The resulting mixture was stirred at 60 °C in an oil bath for 18 h. After cooling to room temperature, the solvent was evaporated under reduced pressure and the crude residue was purified by column chromatography (petroleum ether/EA = 20/1) on silica gel to give product **2f** as a colorless oil (86.7 mg, 97% yield, 65% ee).

General procedure for the removal of TIPS-protecting group of **2r (Scheme 4):** In a test tube, 2-Pic-BH₃ (95 mg, 0.90 mmol) and 10% hydrochloric acid in ethanol (1.0 mL) were added to a solution of **2r** (96.5 mg, 0.30 mmol) in ethanol (1.0 mL) at 0 °C. The reaction mixture was stirred for 4 h at room temperature. The reaction mixture was diluted with saturated Na₂CO₃ solution (3 mL). The aqueous phase was extracted with ethyl acetate (3 × 10 mL) and the combined organic phases were dried over MgSO₄. The solvent was evaporated under reduced pressure and the crude residue was purified by column chromatography (petroleum ether/EA = 4/1) on silica to give product **5** as a colorless oil (44.1 mg, 86% yield).

General procedure for the synthesis of *N*-benzoyl amines for chiral HPLC analysis: In a test tube, triethylamine (20.0 mg, 0.2 mmol) and benzoyl chloride (27.3 mg, 0.12 mmol.) were added to a solution of TIPS-protected hydroxylamine **2** (0.1 mmol) in DCM (1.0 mL) at room temperature. The reaction mixture was stirred at room temperature for 16 h and was then diluted with water. The aqueous phase was extracted with CH₂Cl₂ (3 × 10 mL) and the combined organic phases were dried over Na₂SO₄. The solvent was evaporated under reduced pressure and the crude residue was purified by column chromatography (petroleum ether/EA = 10/1) on silica to give the product for the determination of ee.

J. Mas-Rosello, T. Smejkal and N. Cramer, *Science*, 2020, **368**, 1098.

Characterization of substrates

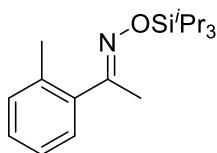


(E)-1-phenylethan-1-one O-triisopropylsilyl oxime (1f): colorless oil. ^1H NMR (300 MHz, CDCl_3 , ppm) δ 7.73-7.64 (m, 2H), 7.38-7.32 (m, 3H), 2.27 (s, 3H), 1.34-1.21 (m, 3H), 1.15-1.07 (m, 18H); ^{13}C NMR (100 MHz, CDCl_3 , ppm) δ 158.5, 137.3, 129.1, 128.5, 126.2, 18.2, 12.2, 12.1

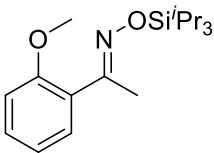
J. Mohr and M. Oestreich, *Angew. Chem. Int. Ed.*, 2014, **53**, 13278-13281.



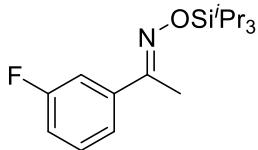
(E)-1-(2-fluorophenyl)ethan-1-one O-triisopropylsilyl oxime (1h): colorless oil. IR (film): 2945, 2867, 1452, 932 cm^{-1} ; ^1H NMR (300 MHz, CDCl_3 , ppm) δ 7.51-7.54 (m, 1H), 7.35-7.27 (m, 1H), 7.16-7.01 (m, 2H), 2.28 (d, $J = 3.0$ Hz, 3H), 1.33-1.20 (m, 3H), 1.15-1.07 (m, 18H); ^{13}C NMR (100 MHz, CDCl_3 , ppm) δ 161.9, 158.2 (d, $J = 224.0$ Hz), 130.2 (d, $J = 8.0$ Hz), 129.6 (d, $J = 4.0$ Hz), 125.8 (d, $J = 12.0$ Hz), 123.9 (d, $J = 3.0$ Hz), 116.1 (d, $J = 22.0$ Hz), 17.9, 14.9 (d, $J = 5.0$ Hz), 12.0; ^{19}F NMR (376 MHz, CDCl_3 , ppm) δ -114.3; HRMS (ESI) calcd. for $\text{C}_{17}\text{H}_{29}\text{FNOSi}$ ($\text{M}+\text{H}$) $^+$: 310.1997, Found: 310.1990.



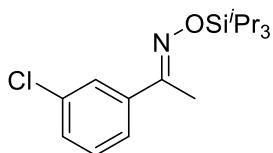
1-(o-tolyl)ethan-1-one O-triisopropylsilyl oxime (1i): colorless oil, a mixture of *Z* and *E* isomers (*Z:E* = 1:5). IR (film): 2944, 2866, 1463, 920 cm^{-1} ; ^1H NMR (300 MHz, CDCl_3 , ppm) δ 7.24-7.16 (m, 4H), 2.35 (s, 2.5H), 2.25 (s, 0.5H), 2.21 (s, 2.5H), 2.12 (s, 0.5H), 1.31-1.19 (m, 3H), 1.15-1.06 (m, 15H), 0.98-0.94 (m, 3H); ^{13}C NMR (100 MHz, CDCl_3 , ppm) δ 161.2, 138.2, 136.0, 130.8, 128.5, 128.4, 125.9, 20.7, 18.2, (18.1 for *Z* isomer), 16.3, 12.2, (12.0 for *Z* isomer); HRMS (ESI) calcd. for $\text{C}_{18}\text{H}_{32}\text{NOSi}$ ($\text{M}+\text{H}$) $^+$: 306.2250, Found: 306.2249.



1-(2-methoxyphenyl)ethan-1-one *O*-triisopropylsilyl oxime (1j): colorless oil, a mixture of *Z* and *E* isomers (*Z:E* = 1:3). IR (film): 2944, 2866, 1463, 920 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.30-7.14 (m, 2H), 6.90-6.77 (m, 2H), 3.74 (s, 2H), 2.14 (s, 2.4H), 2.01 (s, 0.8H), 1.25-1.13 (m, 3H), 1.05-1.03 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 160.6, 157.8, 130.0, 129.8, (128.0 for *Z* isomer), 120.7, 111.4, 55.6, 22.8, 18.3, (18.0 for *Z* isomer), 15.9, 12.2, (12.1 for *Z* isomer); HRMS (ESI) calcd. for C₁₈H₃₂NO₂Si (M+H)⁺: 322.2197, Found: 322.2194.

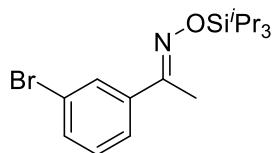


(E)-1-(3-fluorophenyl)ethan-1-one *O*-triisopropylsilyl oxime (1k): colorless oil. IR (film): 2945, 2867, 1463, 878 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.49-7.27 (m, 3H), 7.08-7.00 (m, 1H), 2.26 (s, 3H), 1.34-1.22 (m, 3H), 1.18-1.08 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 163.1 (d, *J* = 243.5 Hz), 157.5 (d, *J* = 2.6 Hz), 139.5 (d, *J* = 7.8 Hz), 129.9 (d, *J* = 8.3 Hz), 121.9 (d, *J* = 2.7 Hz), 115.9 (d, *J* = 21.3 Hz), 113.0 (d, *J* = 23.0 Hz), 18.2, 12.2, 12.0; ¹⁹F NMR (376 MHz, CDCl₃, ppm) δ -113.2; HRMS (ESI) calcd. for C₁₇H₂₉FNOSi (M+H)⁺: 310.1997, Found: 310.1990.

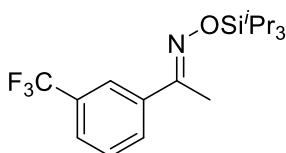


(E)-1-(3-chlorophenyl)ethan-1-one *O*-triisopropylsilyl oxime (1l): colorless oil. IR (film): 2944, 2866, 1461, 990 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.69-7.49 (m, 2H), 7.35-7.21 (m, 2H), 2.25 (s, 3H), 1.36-1.22 (m, 3H), 1.20-1.01 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 157.4, 139.0, 134.5, 129.7, 129.0, 126.3, 124.3, 18.2, 12.1, 12.0; HRMS (ESI) calcd. for C₁₇H₂₉ClNOSi

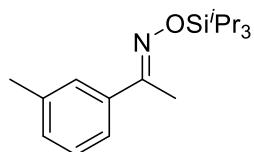
$(M+H)^+$: 326.1702, Found: 326.1693.



(E)-1-(3-bromophenyl)ethan-1-one *O*-triisopropylsilyl oxime (1m): colorless oil. IR (film): 2944, 2866, 1464, 996 cm^{-1} ; ^1H NMR (300 MHz, CDCl_3 , ppm) δ 7.80-7.78 (m, 1H), 7.63-7.59 (m, 1H), 7.49-7.43 (m, 1H), 7.26-7.19 (m, 1H), 2.24 (s, 3H), 1.34-1.22 (m, 3H), 1.21-1.07 (m, 18H). ^{13}C NMR (100 MHz, CDCl_3 , ppm) δ 157.1, 139.1, 131.7, 129.8, 129.0, 124.5, 122.5, 18.0, 11.9, 11.8; HRMS (ESI) calcd. for $\text{C}_{17}\text{H}_{29}\text{BrNOSi}$ ($M+H$) $^+$: 370.1196, Found: 370.1187.

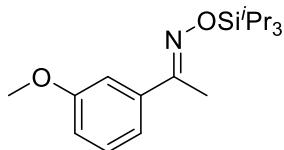


(E)-1-(3-(trifluoromethyl)phenyl)ethan-1-one *O*-triisopropylsilyl oxime (1n): colorless oil. IR (film): 2923, 2867, 1650, 1131 cm^{-1} ; ^1H NMR (300 MHz, CDCl_3 , ppm) δ 7.91-7.86 (m, 2H), 7.61-7.56 (m, 1H), 7.53-7.43 (m, 1H), 2.29 (s, 3H), 1.35-1.22 (m, 3H), 1.21-1.06 (m, 18H); ^{13}C NMR (125 MHz, CDCl_3 , ppm) δ 157.3, 138.0, 131.0 (q, $J = 31.3$ Hz), 129.3, 129.0, 125.7 (q, $J = 3.8$ Hz), 124.3 (q, $J = 271.3$ Hz), 123.0 (q, $J = 3.8$ Hz), 18.2, 12.2, 12.0; ^{19}F NMR (376 MHz, CDCl_3 , ppm) δ -62.8; HRMS (ESI) calcd. for $\text{C}_{18}\text{H}_{29}\text{F}_3\text{NOSi}$ ($M+H$) $^+$: 360.1965, Found: 360.1956.

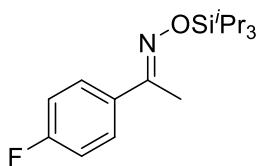


(E)-1-(m-tolyl)ethan-1-one *O*-triisopropylsilyl oxime (1o): colorless oil. IR (film): 2944, 2866, 1463, 939 cm^{-1} ; ^1H NMR (300 MHz, CDCl_3 , ppm) δ 7.50-7.44 (m, 2H), 7.27-7.11 (m, 2H), 2.37 (s, 3H), 2.26 (s, 3H), 1.34-1.21 (m, 3H), 1.21-1.06 (m, 18H); ^{13}C NMR (100 MHz, CDCl_3 , ppm) δ

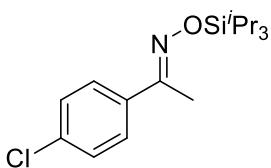
158.6, 138.0, 137.3, 129.8, 128.4, 126.9, 123.4, 21.8, 18.2, 12.2; HRMS (ESI) calcd. for C₁₈H₃₂NOSi (M+H)⁺: 306.2250, Found: 306.2248.



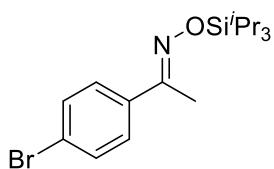
(E)-1-(3-methoxyphenyl)ethan-1-one O-triisopropylsilyl oxime (1p): colorless oil. IR (film): 2944, 2866, 1464, 942 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.33-7.23 (m, 3H), 6.93-6.87 (m, 1H), 3.82 (s, 3H), 2.26 (s, 3H), 1.34-1.20 (m, 3H), 1.20-1.06 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 159.7, 158.3, 138.7, 129.4, 118.8, 114.8, 111.6, 55.4, 18.2, 12.2, 12.1; HRMS (ESI) calcd. for C₁₈H₃₂NO₂Si (M+H)⁺: 322.2197, Found: 322.2197.



(E)-1-(4-fluorophenyl)ethan-1-one O-triisopropylsilyl oxime (1q): colorless oil. IR (film): 2945, 2867, 1511, 930 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.69-7.62 (m, 2H), 7.08-7.00 (m, 2H), 2.25 (s, 3H), 1.33-1.21 (m, 3H), 1.16-1.07 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 164.7, 162.2, 157.5, 133.4, 128.0 (d, *J* = 8.0 Hz), 115.4 (d, *J* = 21.0 Hz), 18.2, 12.2, 12.0; ¹⁹F NMR (376 MHz, CDCl₃, ppm) δ -112.9; HRMS (ESI) calcd. for C₁₇H₂₉FNOSi (M+H)⁺: 310.1997, Found: 310.1991.

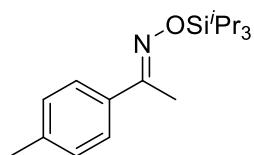


(E)-1-(4-chlorophenyl)ethan-1-one *O*-triisopropylsilyl oxime (1r): colorless oil. IR (film): 2944, 2866, 1461, 990 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.61 (d, *J* = 8.6 Hz, 2H), 7.32 (d, *J* = 8.6 Hz, 2H), 2.25 (s, 3H), 1.33-1.21 (m, 3H), 1.16-1.06 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 157.5, 135.7, 135.1, 128.7, 127.4, 18.2, 12.2, 12.0; HRMS (ESI) calcd. for C₁₇H₂₉ClNOSi (M+H)⁺: 326.1702, Found: 326.1693.

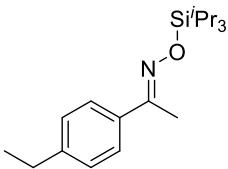


(E)-1-(4-bromophenyl)ethan-1-one *O*-triisopropylsilyl oxime (1s): colorless oil; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.55 (d, *J* = 9.0 Hz, 2H), 7.48 (d, *J* = 9.0 Hz, 2H), 2.24 (s, 3H), 1.33-1.21 (m, 3H), 1.16-1.06 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 157.6, 136.2, 131.6, 127.7, 123.3, 18.2, 12.1, 11.9.

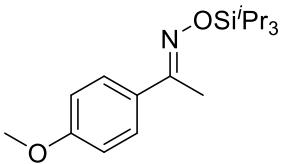
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(E)-1-(p-tolyl)ethan-1-one *O*-triisopropylsilyl oxime (1t): colorless oil. IR (film): 2944, 2866, 1463, 926 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.58 (d, *J* = 8.1 Hz, 2H), 7.16 (d, *J* = 8.1 Hz, 2H), 2.35 (s, 3H), 2.25 (s, 3H), 1.33-1.21 (m, 3H), 1.16-1.06 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 158.4, 139.0, 134.5, 129.2, 126.1, 21.5, 18.2, 12.2, 12.0; HRMS (ESI) calcd. for C₁₈H₃₂NOSi (M+H)⁺: 306.2250, Found: 306.2247.

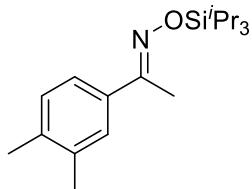


(E)-1-(4-ethylphenyl)ethan-1-one O-triisopropylsilyl oxime (1u): colorless oil. IR (film): 2943, 2866, 1462, 925 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.62 (d, *J* = 8.4 Hz, 2H), 7.19 (d, *J* = 8.4 Hz, 2H), 2.65 (q, *J* = 7.7 Hz, 2H), 2.25 (s, 3H), 1.33-1.21 (m, 3H), 1.24 (t, *J* = 7.5 Hz, 3H), 1.14-1.08 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 161.9, 136.6, 128.9, 128.5, 126.4, 28.0, 20.3, 18.2, 14.6, 12.2; HRMS (ESI) calcd. for C₁₉H₃₄NOSi (M+H)⁺: 320.2404, Found: 320.2403.



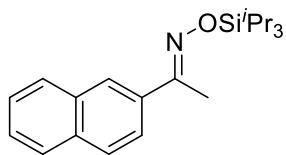
(E)-1-(4-methoxyphenyl)ethan-1-one O-triisopropylsilyl oxime (1v): colorless oil. ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.62 (d, *J* = 8.8 Hz, 2H), 6.87 (d, *J* = 8.7 Hz, 2H), 3.79 (s, 3H), 2.24 (s, 3H), 1.33-1.21 (m, 3H), 1.16-1.05 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 160.5, 157.9, 129.9, 127.5, 113.9, 55.4, 18.2, 12.2, 11.9.

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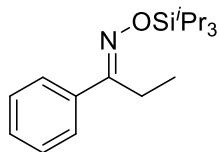


(E)-1-(3,4-dimethylphenyl)ethan-1-one O-triisopropylsilyl oxime (1w): colorless oil. IR (film): 2943, 2866, 1463, 880 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.46-7.38 (m, 2H), 7.11 (d, *J* = 7.8 Hz, 1H), 2.28 (s, 3H), 2.26 (s, 3H), 2.24 (s, 3H), 1.33-1.21 (m, 3H), 1.17-1.06 (m, 18H); ¹³C NMR

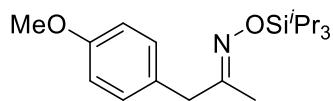
(100 MHz, CDCl₃, ppm) δ 158.3, 137.5, 136.5, 134.7, 129.5, 127.4, 123.5, 20.1, 19.7, 18.1, 12.1, 12.0; HRMS (ESI) calcd. for C₁₉H₃₄NOSi (M+H)⁺: 320.2404, Found: 320.2405.



(E)-1-(naphthalen-2-yl)ethan-1-one O-triisopropylsilyl oxime (1x): colorless oil. IR (film): 2944, 2866, 1463, 925 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 8.02-7.93 (m, 2H), 7.89-7.77 (m, 3H), 7.51-7.43 (m, 2H), 2.39 (s, 3H), 1.39-1.25 (m, 3H), 1.18-1.08 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 158.4, 134.7, 133.8, 133.3, 128.6, 128.0, 127.8, 126.6, 126.4, 125.9, 123.7, 18.2, 12.2, 11.9; HRMS (ESI) calcd. for C₂₁H₃₂NOSi (M+H)⁺: 342.2248, Found: 342.2249.



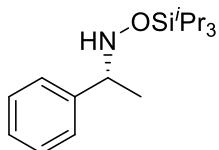
(E)-1-phenylpropan-1-one O-triisopropylsilyl oxime (1y): colorless oil. IR (film): 2943, 2866, 1463, 934 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.70-7.62 (m, 2H), 7.40-7.31 (m, 3H), 2.81 (q, J = 7.6 Hz, 2H), 1.34-1.23 (m, 3H), 1.22-1.11 (m, 2H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 163.0, 136.3, 129.0, 128.6, 126.4, 19.6, 18.2, 12.2, 11.2; HRMS (ESI) calcd. for C₁₈H₃₂NOSi (M+H)⁺: 306.2248, Found: 306.2243.



(E)-1-(4-methoxyphenyl)propan-2-one O-triisopropylsilyl oxime (1z): colorless oil. IR (film): 2942, 2864, 1463, 927 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.54 (d, J = 5.6 Hz, 2H), 6.79 (d, J = 5.6 Hz, 2H), 3.71 (s, 3H), 3.22 (s, 2H), 2.16 (s, 3H), 1.27-1.11 (m, 3H), 1.06-0.95 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 160.2, 157.6, 129.6, 127.2, 113.5, 55.1, 45.8, 17.9, 11.9, 11.6;

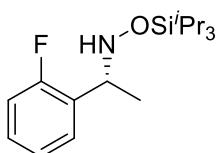
HRMS (ESI) calcd. for $C_{19}H_{34}NO_2Si$ ($M+H$)⁺: 336.2359, Found: 336.2353.

Characterization of products:

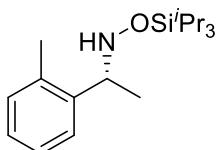


(R)-N-(1-phenylethyl)-O-(triisopropylsilyl)hydroxylamine (2f): colorless oil, 86.7 mg, 97% yield, 65% ee, $[\alpha]_D^{24} = +22.6$ (*c* 0.50, CHCl₃). ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.37-7.22 (m, 5H), 4.15 (q, *J* = 6.7 Hz, 1H), 1.47 (d, *J* = 6.7 Hz, 3H), 1.18-1.08 (m, 3H), 1.08-0.96 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 141.9, 128.6, 127.9, 127.6, 62.6, 19.0, 18.3, 12.0.

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(R)-N-(1-(2-fluorophenyl)ethyl)-O-(triisopropylsilyl)hydroxylamine (2h): colorless oil, 90.1 mg, 96% yield, 45% ee, $[\alpha]_D^{24} = +14.2$ (*c* 0.50, CHCl₃). IR (film): 2945, 2868, 1492, 1231 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.28 (m, 1H), 7.27-7.17 (m, 1H), 7.14-7.06 (m, 1H), 7.06-7.69 (m, 1H), 4.40 (q, *J* = 6.7 Hz, 1H), 1.41 (d, *J* = 6.8 Hz, 3H), 1.19-1.07 (m, 3H), 1.07-0.99 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 161.2 (d, *J* = 244.0 Hz), 130.2 (d, *J* = 13.4 Hz), 128.7 (d, *J* = 12.0 Hz), 128.7, 124.2 (d, *J* = 3.4 Hz), 115.6 (d, *J* = 22.4 Hz), 56.5, 18.7, 18.4, 12.0; ¹⁹F NMR (376 MHz, CDCl₃, ppm) δ -113.6; HRMS (ESI) calcd. for $C_{17}H_{31}FNOSi$ ($M+H$)⁺: 312.2153, Found: 312.2154.

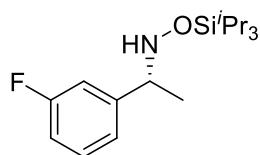


(R)-N-(1-(o-tolyl)ethyl)-O-(triisopropylsilyl)hydroxylamine (2i): colorless oil, 85.2 mg, 92%

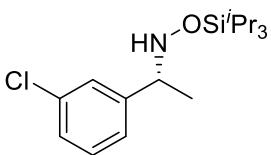
yield, 52% ee, $[\alpha]_D^{24} = +15.1$ (c 1.00, CHCl₃). IR (film): 2943, 2866, 1465, 1090 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.34-7.30 (m, 1H), 7.21-7.11 (m, 3H), 5.00 (s, 1H), 4.38 (q, $J = 6.6$ Hz, 1H), 2.39 (s, 3H), 1.39 (d, $J = 6.6$ Hz, 3H), 1.19-1.12 (m, 3H), 1.12-1.02 (m, 18H); ¹³C NMR (100 MHz, CDCl₃) δ 141.3, 136.1, 130.5, 127.2, 126.3, 126.0, 58.1, 19.6, 18.4, 18.0, 12.1; HRMS (ESI) calcd. for C₁₈H₃₄NOSi (M+H)⁺: 308.2404, Found: 308.2406.



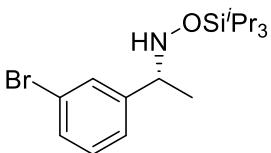
(R)-N-(1-(1-methoxyphenyl)ethyl)-O-(triisopropylsilyl)hydroxylamine (2j): colorless oil, 81.2 mg, 88% yield, 48% ee, $[\alpha]_D^{24} = +17.7$ (c 0.50, CHCl₃). IR (film): 2906, 2361, 1634, 1461, 1245 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.26-7.21 (m, 1H), 6.95-6.77 (m, 3H), 5.19 (br s, 1H), 4.10 (q, $J = 6.6$ Hz, 1H), 3.80 (s, 3H), 1.45 (d, $J = 6.6$ Hz, 3H), 1.18-1.09 (m, 3H), 1.09-1.01 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 159.9, 129.6, 119.9, 113.4, 113.2, 62.6, 55.4, 18.9, 18.3, 12.0; HRMS (ESI) calcd. for C₁₈H₃₄NO₂Si (M+H)⁺: 324.2353, Found: 324.2353.



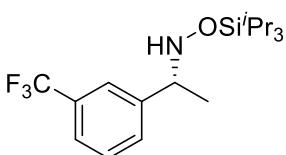
(R)-N-(1-(3-fluorophenyl)ethyl)-O-(triisopropylsilyl)hydroxylamine (2k): colorless oil, 86.9 mg, 93% yield, 61% ee, $[\alpha]_D^{24} = +18.2$ (c 0.51, CHCl₃). IR (film): 2946, 2869, 1592, 1317 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.33-7.22 (m, 1H), 7.14-6.89 (m, 3H), 4.10 (q, $J = 6.5$ Hz, 1H), 1.40 (d, $J = 6.7$ Hz, 3H), 1.20-1.09 (m, 3H), 1.09-0.99 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 163.1 (d, $J = 243.8$ Hz), 145.8 (d, $J = 6.8$ Hz), 129.8 (d, $J = 8.2$ Hz), 123.2 (d, $J = 2.5$ Hz), 114.5 (d, $J = 5.8$ Hz), 114.3 (d, $J = 5.6$ Hz), 62.0, 19.6, 18.4, 12.0; ¹⁹F NMR (376 MHz, CDCl₃, ppm) δ -112.3; HRMS (ESI) calcd. for C₁₇H₃₁FNOSi (M+H)⁺: 312.2153, Found: 312.2153.



(R)-N-(1-(3-chlorophenyl)ethyl)-O-(triisopropylsilyl)hydroxylamine (2l): colorless oil, 86.1 mg, 86% yield, 60% ee, $[\alpha]_D^{24} = +20.6$ (c 0.50, CHCl₃). IR (film): 2943, 2866, 1465 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.31 (s, 1H), 7.25-7.19 (m, 3H), 4.06 (q, $J = 6.8$ Hz, 1H), 1.38 (d, $J = 6.7$ Hz, 3H), 1.19-1.08 (m, 3H), 1.08-0.99 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 145.2, 134.3, 129.7, 127.8, 127.6, 125.8, 62.0, 19.5, 18.4, 12.0; HRMS (ESI) calcd. for C₁₇H₃₁ClNO₂Si (M+H)⁺: 328.1858, Found: 328.1857.

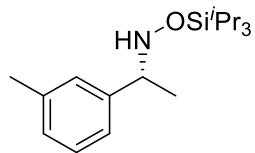


(R)-N-(1-(3-bromophenyl)ethyl)-O-(triisopropylsilyl)hydroxylamine (2m): colorless oil, 100.3 mg, 89% yield, 62% ee, $[\alpha]_D^{24} = +22.1$ (c 0.51, CHCl₃). IR (film): 2943, 2866, 1461 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.50 (s, 2H), 7.28-7.17 (m, 2H), 4.07 (q, $J = 6.8$ Hz, 1H), 1.40 (d, $J = 6.7$ Hz, 3H), 1.21-1.10 (m, 3H), 1.10-1.00 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 145.5, 130.7, 130.6, 130.0, 126.3, 122.5, 62.0, 19.5, 18.4, 18.3, 12.0; HRMS (ESI) calcd. for C₁₇H₃₁BrNO₂Si (M+H)⁺: 372.1353, Found: 372.1353.

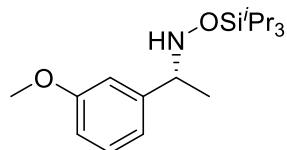


(R)-N-(1-(3-(trifluoromethyl)phenyl)ethyl)-O-(triisopropylsilyl)hydroxylamine (2n): colorless oil, 99.3 mg, 90% yield, 68% ee, $[\alpha]_D^{24} = +23.3$ (c 0.50, CHCl₃). IR (film): 2881, 2362, 1328, 1129 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.63-7.37 (m, 4H), 4.15 (q, $J = 6.9$ Hz, 1H), 1.42 (d, $J = 6.6$ Hz, 3H), 1.17-1.08 (m, 3H), 1.08-0.96 (m, 18H); ¹³C NMR (125 MHz, CDCl₃, ppm) δ 144.2,

131.0, 130.9 (q, $J = 31.8$ Hz), 128.8, 124.5 (q, $J = 270.3$ Hz), 124.5 (q, $J = 3.7$ Hz), 124.3 (q, $J = 3.6$ Hz), 62.0, 19.4, 18.3, 12.0; ^{19}F NMR (376 MHz, CDCl_3 , ppm) δ -62.4; HRMS (ESI) calcd. for $\text{C}_{18}\text{H}_{31}\text{F}_3\text{NOSi} (\text{M}+\text{H})^+$: 362.2122, Found: 362.2120.



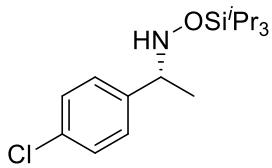
(R)-N-(1-(m-tolyl)ethyl)-O-(triisopropylsilyl)hydroxylamine (2o): colorless oil, 98.5 mg, 99% yield, 60% ee, $[\alpha]_D^{24} = +19.7$ (c 0.50, CHCl_3). IR (film): 2943, 2866, 1464 cm^{-1} ; ^1H NMR (300 MHz, CDCl_3 , ppm) δ 7.24-7.03 (m, 4H), 4.06 (q, $J = 6.6$ Hz, 1H), 2.34 (s, 3H), 1.41 (d, $J = 6.6$ Hz, 3H), 1.20-1.10 (m, 3H), 1.10-1.00 (m, 18H); ^{13}C NMR (100 MHz, CDCl_3 , ppm) δ 142.7, 138.7, 128.2, 128.1, 124.4, 62.2, 21.5, 19.4, 18.2, 11.9; HRMS (ESI) calcd. for $\text{C}_{18}\text{H}_{34}\text{NOSi} (\text{M}+\text{H})^+$: 308.2404, Found: 308.2406.



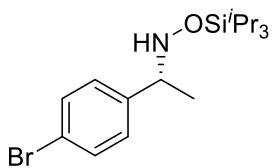
(R)-N-(1-(3-methoxyphenyl)ethyl)-O-(triisopropylsilyl)hydroxylamine (2p): colorless oil, 86.6 mg, 84% yield, 57% ee, $[\alpha]_D^{24} = +18.6$ (c 0.51, CHCl_3). IR (film): 2904, 2361, 1604, 1041 cm^{-1} ; ^1H NMR (300 MHz, CDCl_3 , ppm) δ 7.25-7.19 (m, 1H), 6.92-6.77 (m, 3H), 4.08 (q, $J = 6.6$ Hz, 1H), 3.80 (s, 3H), 1.42 (d, $J = 6.8$ Hz, 3H), 1.19-1.11 (m, 3H), 1.11-1.01 (m, 18H); ^{13}C NMR (100 MHz, CDCl_3 , ppm) δ 159.6, 143.7, 129.3, 119.7, 113.2, 112.9, 62.3, 55.2, 19.0, 18.1, 11.8; HRMS (ESI) calcd. for $\text{C}_{18}\text{H}_{34}\text{NO}_2\text{Si} (\text{M}+\text{H})^+$: 324.2353, Found: 324.2358.



(R)-N-(1-(4-fluorophenyl)ethyl)-O-(triisopropylsilyl)hydroxylamine (2q): colorless oil, 85.2 mg, 91% yield, 55% ee, $[\alpha]_D^{24} = +16.3$ (*c* 0.50, CHCl₃). IR (film): 2973, 2872, 1605, 1228 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.34-7.22 (m, 2H), 7.05-6.94 (m, 2H), 4.89 (s, 1H), 4.07 (q, *J* = 5.6 Hz, 1H), 1.39 (d, *J* = 6.6 Hz, 3H), 1.17-1.07 (m, 3H), 1.09-0.97 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 162.3 (d, *J* = 243.4 Hz), 138.7 (d, *J* = 3.0 Hz), 129.1 (d, *J* = 7.9 Hz), 115.2 (d, *J* = 21.0 Hz), 61.7, 19.6, 18.4, 12.0; ¹⁹F NMR (376 MHz, CDCl₃, ppm) δ -111.9; HRMS (ESI) calcd. for C₁₇H₃₁FNOSi (M+H)⁺: 312.2153, Found: 312.2155.



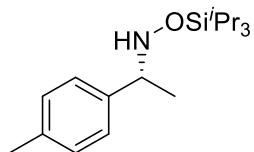
(R)-N-(1-(4-chlorophenyl)ethyl)-O-(triisopropylsilyl)hydroxylamine (2r): colorless oil, 92.1 mg, 92% yield, 56% ee, $[\alpha]_D^{24} = +21.8$ (*c* 0.46, CHCl₃). IR (film): 2944, 2867, 1464, 1087 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.34-7.21 (m, 4H), 4.07 (q, *J* = 6.5 Hz, 1H), 1.38 (d, *J* = 6.7 Hz, 3H), 1.20-1.08 (m, 3H), 1.08-0.98 (m, 18H); ¹³C NMR (125 MHz, CDCl₃, ppm) δ 141.5, 133.1, 128.9, 128.5, 61.7, 19.5, 18.3, 12.0; HRMS (ESI) calcd. for C₁₇H₃₁ClNOSi (M+H)⁺: 328.1858, Found: 328.1856.



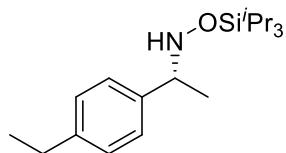
(R)-N-(1-(4-bromophenyl)ethyl)-O-(triisopropylsilyl)hydroxylamine (2s): colorless oil, 106.1 mg, 89% yield, 59% ee, $[\alpha]_D^{24} = +21.8$ (*c* 0.52, CHCl₃). ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.44 (d, *J* = 8.3 Hz, 2H), 7.20 (d, *J* = 8.3 Hz, 3H), 4.05 (q, *J* = 6.8 Hz, 1H), 1.38 (d, *J* = 6.7 Hz, 3H), 1.19-1.08 (m, 3H), 1.08-0.99 (m, 18H); ¹³C NMR (125 MHz, CDCl₃, ppm) δ 142.1, 131.5, 129.3,

121.3, 61.8, 19.5, 18.4, 18.4, 12.0.

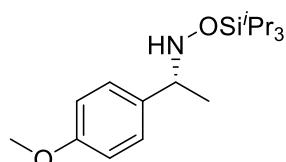
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(R)-N-(1-(p-tolyl)ethyl)-O-(triisopropylsilyl)hydroxylamine (2t): colorless oil, 84.2 mg, 89% yield, 54% ee, $[\alpha]_D^{24} = +18.7$ (*c* 0.53, CHCl₃). IR (film): 2943, 2866, 1465, 1090 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.29-7.08 (m, 4H), 4.06 (q, *J* = 6.3 Hz, 1H), 2.33 (s, 3H), 1.41 (d, *J* = 6.5 Hz, 3H), 1.19-1.11 (m, 3H), 1.11-1.00 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 139.9, 137.2, 129.2, 127.5, 62.1, 21.3, 19.6, 18.4, 12.1; HRMS (ESI) calcd. for C₁₈H₃₄NOSi (M+H)⁺: 308.2404, Found: 308.2406.



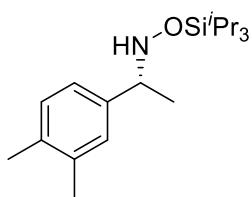
(R)-N-(1-(4-ethylphenyl)ethyl)-O-(triisopropylsilyl)hydroxylamine (2u): colorless oil, 91.1 mg, 92% yield, 54% ee, $[\alpha]_D^{24} = +18.9$ (*c* 0.58, CHCl₃). IR (film): 2943, 2866, 1462, 992 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.28-7.21 (m, 2H), 7.15 (d, *J* = 7.9 Hz, 2H), 4.08 (q, *J* = 6.4 Hz, 1H), 2.63 (q, *J* = 7.5 Hz, 2H), 1.43 (d, *J* = 6.5 Hz, 3H), 1.23 (t, *J* = 7.6 Hz, 3H), 1.20-1.09 (m, 3H), 1.09-1.00 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 158.4, 145.4, 134.8, 128.0, 126.2, 28.9, 18.2, 15.7, 12.2, 12.0; HRMS (ESI) calcd. for C₁₉H₃₆NOSi (M+H)⁺: 322.2561, Found: 322.2562.



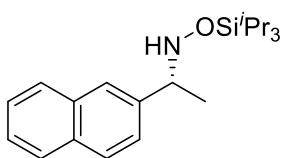
(R)-N-(1-(4-methoxyphenyl)ethyl)-O-(triisopropylsilyl)hydroxylamine (2v): colorless oil, 84.4

mg, 89% yield, 51% ee, $[\alpha]_D^{24} = +17.0$ (c 0.50, CHCl_3). ^1H NMR (300 MHz, CDCl_3 , ppm) δ 7.25 (d, $J = 8.6$ Hz, 2H), 6.85 (d, $J = 8.7$ Hz, 3H), 4.87 (s, 1H), 4.05 (q, $J = 6.6$ Hz, 1H), 3.79 (s, 3H), 1.40 (d, $J = 6.6$ Hz, 3H), 1.18-1.11 (m, 3H), 1.10-1.00 (m, 18H); ^{13}C NMR (100 MHz, CDCl_3 , ppm) δ 159.1, 134.9, 128.7, 113.8, 61.7, 55.4, 19.5, 18.4, 12.1.

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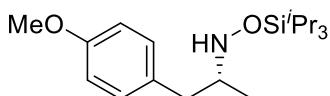
(R)-N-(1-(3,4-dimethylphenyl)ethyl)-O-(triisopropylsilyl)hydroxylamine (2w): colorless oil, 91.7 mg, 94% yield, 58% ee, $[\alpha]_D^{24} = +19.2$ (c 0.50, CHCl_3). IR (film): 2925, 2865, 1635, 1462, 1085 cm^{-1} ; ^1H NMR (300 MHz, CDCl_3 , ppm) δ 7.13-7.01 (m, 3H), 4.88 (s, 1H), 4.04 (q, $J = 6.4$ Hz, 1H), 2.25 (s, 3H), 2.24 (s, 3H), 1.40 (d, $J = 6.5$ Hz, 3H), 1.22-1.12 (m, 3H), 1.11-1.00 (m, 18H); ^{13}C NMR (100 MHz, CDCl_3 , ppm) δ 140.3, 136.6, 135.8, 129.7, 128.9, 124.9, 62.1, 20.0, 19.6, 18.4, 12.0; HRMS (ESI) calcd. for $\text{C}_{19}\text{H}_{36}\text{NOSi} (\text{M}+\text{H})^+$: 322.2561, Found: 322.2566.



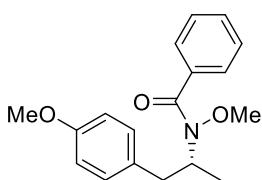
(R)-N-(1-(naphthalen-2-yl)ethyl)-O-(triisopropylsilyl)hydroxylamine (2x): colorless oil, 92.9 mg, 92% yield, 64% ee, $[\alpha]_D^{24} = +21.8$ (c 0.55, CHCl_3). IR (film): 2943, 2865, 1463, 996 cm^{-1} ; ^1H NMR (300 MHz, CDCl_3 , ppm) δ 7.84-7.72 (m, 4H), 7.50-7.42 (m, 3H), 4.27 (q, $J = 6.6$ Hz, 1H), 1.51 (d, $J = 6.6$ Hz, 3H), 1.20-1.10 (m, 3H), 1.11-1.01 (m, 18H); ^{13}C NMR (100 MHz, CDCl_3 , ppm) δ 140.4, 133.5, 133.1, 128.0, 128.0, 127.8, 126.2, 126.1, 125.9, 125.8, 62.5, 19.6, 18.4, 12.0; HRMS (ESI) calcd. for $\text{C}_{21}\text{H}_{34}\text{NOSi} (\text{M}+\text{H})^+$: 344.2407, Found: 344.2409.



(R)-N-(1-phenylpropyl)-O-(triisopropylsilyl)hydroxylamine (2y): colorless oil, 86.3 mg, 90% yield, 60% ee, $[\alpha]_D^{24} = +19.9$. (c 0.53, CHCl₃). IR (film): 2943, 2866, 1463 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.34-7.23 (m, 5H), 3.95-3.87 (m, 1H), 1.97-1.82 (m, 1H), 1.70-1.51 (m, 1H), 1.31-1.10 (m, 3H), 1.09-0.97 (m, 18H), 0.88 (t, $J = 7.3$ Hz, 3H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 141.8, 128.4, 128.1, 127.5, 69.4, 26.4, 18.4, 12.0, 10.9; HRMS (ESI) calcd. for C₁₈H₃₄NOSi (M+H)⁺: 308.2404, Found: 308.2404.



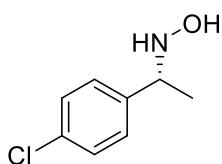
(R)-N-(1-(4-methoxyphenyl)propan-2-yl)-O-(triisopropylsilyl)hydroxylamine (2z): colorless oil, 88.3 mg, 87% yield, 33% ee, $[\alpha]_D^{24} = -11.9$. (c 0.54, CHCl₃). IR (film): 2943, 2865, 1463 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.07-7.01 (m, 2H), 6.64 (d, $J = 6.6$ Hz, 2H), 4.66 (s, 1H), 3.84 (m, 1H), 3.58 (s, 3H), 2.96 (dd, $J = 13.8, 9.0$ Hz, 1H), 2.57 (dd, $J = 13.8, 5.7$ Hz, 1H), 1.19 (d, $J = 6.3$ Hz, 3H), 0.97-0.94 (m, 3H), 0.94-0.80 (m, 18H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 158.4, 134.4, 128.1, 113.3, 64.0, 55.3, 19.0, 17.9, 11.5; HRMS (ESI) calcd. for C₁₉H₃₆NO₂Si (M+H)⁺: 338.2515, Found: 338.2509.



(R)-N-methoxy-N-(1-(4-methoxyphenyl)propan-2-yl)benzamide: According to the general procedure, (R)-**2z** was converted to the *N*-benzoyl derivative for chiral HPLC analysis. ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.40-7.21 (m, 7H), 7.02 (d, $J = 8.4$ Hz, 2H), 6.82-6.76 (m, 2H), 4.34 (s,

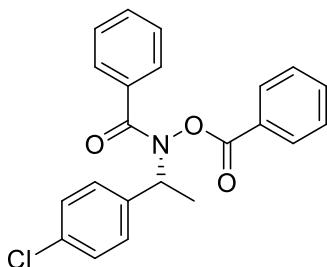
1H), 3.77 (s, 3H), 3.70 (s, 3H), 3.03 (dd, $J = 13.8, 9.0$ Hz, 1H), 2.64 (dd, $J = 13.8, 5.7$ Hz, 1H), 1.36 (d, $J = 6.3$ Hz, 3H); ^{13}C NMR (100 MHz, CDCl_3 , ppm) δ 170.4, 158.4, 135.2, 130.5, 130.3, 130.2, 128.0, 127.4, 113.8, 64.0, 55.3, 39.2, 18.3.

J. Mas-Rosello, T. Smejkal and N. Cramer, *Science*, 2020, **368**, 1098.

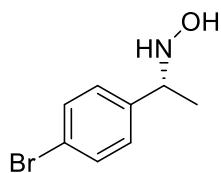


(R)-N-(1-(4-chlorophenyl)ethyl)hydroxylamine (5): white solid, 44.1 mg, 86% yield, 55% ee, $[\alpha]_D^{25} = +8.7$. (c 0.52, CHCl_2) ($[\alpha]_D^{22} = -47.0$ (c 2.0, CHCl_2) for *S*-isomer)^{lit}; ^1H NMR (300 MHz, CDCl_3 , ppm) δ 7.36-7.27 (m, 4H), 4.12 (q, $J = 6.7$ Hz, 1H), 1.33 (d, $J = 6.7$ Hz, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 141.4, 133.4, 128.9, 128.7, 61.4, 19.8.

D. A. Tickell, M. F. Mahon, S. D. Bull and T. D. James, *Org. Lett.*, 2013, **15**, 860-863.

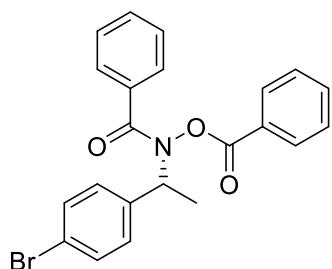


(R)-N-(benzoyloxy)-N-(1-(4-chlorophenyl)ethyl)benzamide: According to the general procedure, **5** was converted to the *N*-benzoyl derivative for chiral HPLC analysis. colorless oil, 26.3 mg, 69% yield, 55% ee. IR (film): 2987, 2874, 1724, 1137 cm^{-1} ; ^1H NMR (300 MHz, CDCl_3 , ppm) δ 7.83-7.73 (m, 2H), 7.58-7.46 (m, 3H), 7.39-7.21 (m, 9H), 5.70 (q, $J = 6.9$ Hz, 1H), 1.60 (d, $J = 6.9$ Hz, 3H); ^{13}C NMR (100 MHz, CDCl_3 , ppm) δ 170.4, 168.0, 134.0, 133.9, 133.8, 130.9, 129.7, 128.7, 128.7, 128.3, 127.5, 126.8, 120.5, 57.8, 29.7, 17.3; HRMS (ESI) calcd. for $\text{C}_{22}\text{H}_{18}\text{O}_3\text{NClNa}$ ($\text{M}+\text{Na}$)⁺: 402.0867, Found: 402.0871.



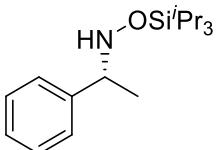
(R)-N-(1-(4-bromophenyl)ethyl)hydroxylamine (6): white solid, 56.7 mg, 88% yield, 61% ee, $[\alpha]_D^{25} = +9.6$. (c 0.60, CHCl₃). ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.47 (d, $J = 8.4$ Hz, 2H), 7.22 (d, $J = 8.3$ Hz, 2H), 4.10 (q, $J = 6.7$ Hz, 1H), 1.34 (d, $J = 6.7$ Hz, 3H); ¹³C NMR (100 MHz, CDCl₃) δ 141.8, 131.8, 129.1, 121.5, 61.4, 19.7.

G. Zeng, H. Li, Y. Wei, W. Xuan, R. Zhang, L. E. Breden, W. Wang and F.-S. Liang, *ACS Synth. Biol.*, 2017, **6**, 921–927.



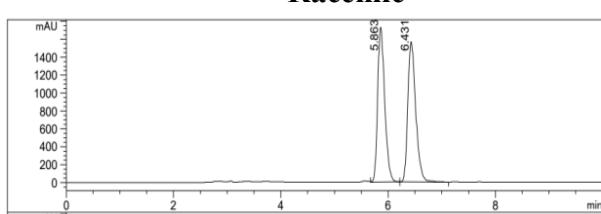
N-(benzoyloxy)-N-(1-(4-bromophenyl)ethyl)benzamide: According to the general procedure, **6** was converted to the *N*-benzoyl derivative for chiral HPLC analysis. colorless oil; 24.3 mg, 57% yield, 61% ee. IR (film): 2987, 2874, 1725, 1139 cm⁻¹; ¹H NMR (300 MHz, CDCl₃, ppm) δ 7.92-7.80 (m, 2H), 7.67-7.59 (m, 3H), 7.53-7.27 (m, 9H), 5.75 (q, $J = 6.7$ Hz, 1H), 1.67 (d, $J = 6.9$ Hz, 3H); ¹³C NMR (100 MHz, CDCl₃, ppm) δ 173.6, 171.2, 134.0, 133.9, 131.7, 131.0, 129.7, 128.9, 128.7, 128.4, 128.3, 127.5, 126.8, 121.9, 57.9, 17.3; HRMS (ESI) calcd. for C₂₂H₁₈O₃NBrNa (M+Na)⁺: 446.0362, Found: 446.0366.

The chromatography for the determination of enantiomeric excess

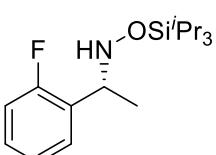
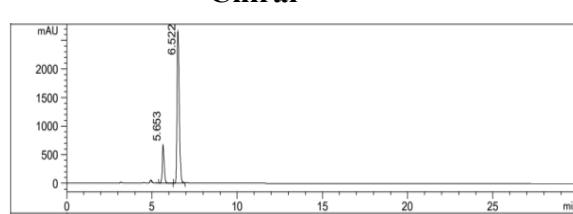


HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (97/3); flow rate: 1.0 mL/min; detection: UV 210 nm

Racemic

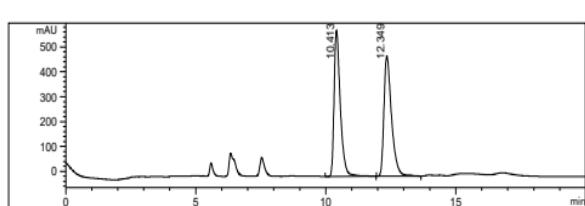


Chiral

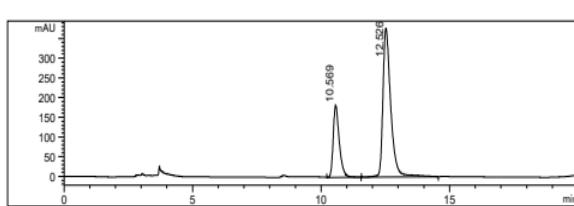


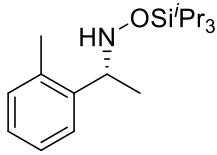
HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (97/3); flow rate: 1.0 mL/min; detection: UV 210 nm

Racemic



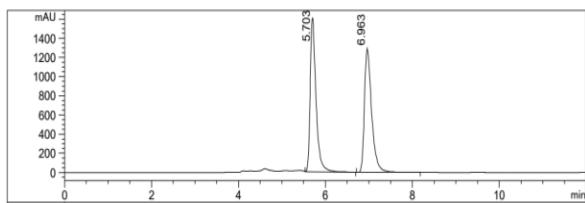
Chiral



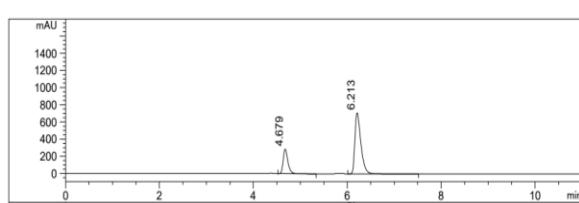


HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (95/5); flow rate: 1.0 mL/min; detection: UV 210 nm

Racemic

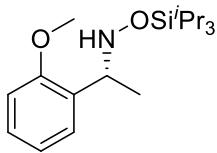


Chiral



Peak	RT	Area	Area %
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1	5.703	1.493e4	50.062
2	6.963	1.490e4	49.938

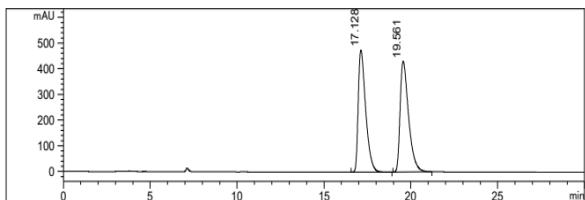
Peak	RT	Area	Area %
#	[min]		
1	4.679	1.999e3	24.113
2	6.213	6.290e3	75.887



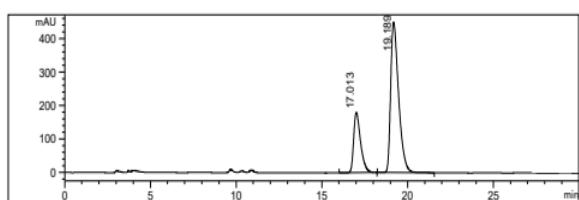
2j

HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (92/8); flow rate: 1.0 mL/min; detection: UV 210 nm

Racemic

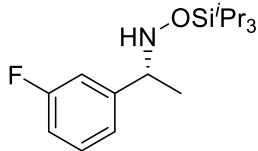


Chiral



Peak	RT	Area	Area %
#	[min]		
1	17.128	1.420e4	49.231
2	19.561	1.464e4	50.769

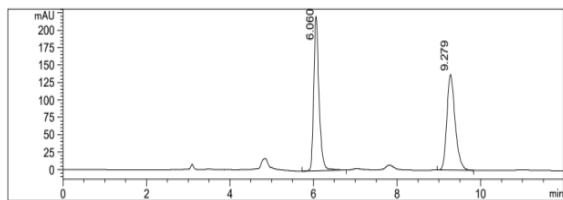
Peak	RT	Area	Area %
#	[min]		
1	17.013	5.076e3	25.884
2	19.189	1.453e4	74.116



2k

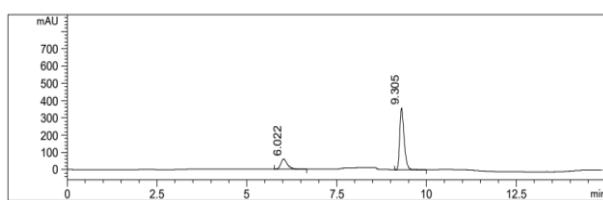
HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (96/4); flow rate: 1.0 mL/min; detection: UV 210 nm

Racemic

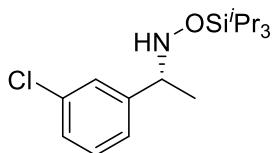


Peak	RT	Area	Area %
#	[min]	-----	-----
1	6.060	1.947e3	50.782
2	9.279	1.887e3	49.218

Chiral



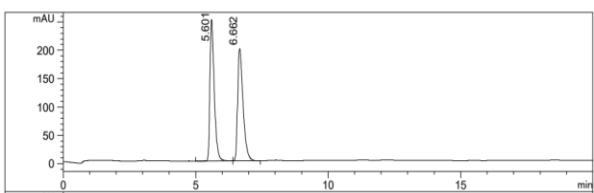
Peak	RT	Area	Area %
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1	6.022	737.954	19.523
2	9.305	3.042e3	80.477



2l

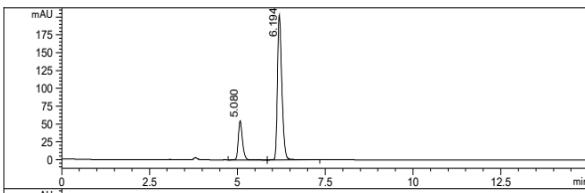
HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (97/3); flow rate: 1.0 mL/min; detection: UV 210 nm

Racemic

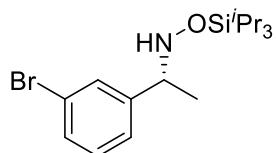


Peak	RT	Area	Area %
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1	5.601	2.841e3	50.323
2	6.662	2.805e3	49.677

Chiral

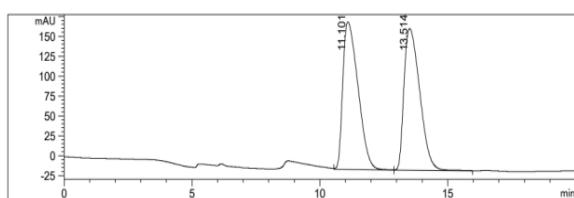


Peak	RT	Area	Area %
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1	5.080	445.616	20.061
2	6.194	1.776e3	79.939



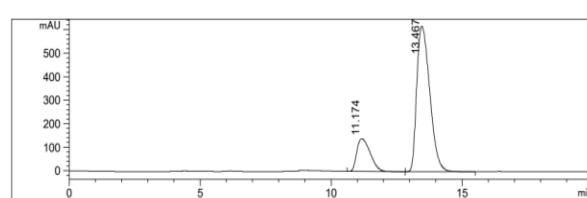
HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (97/3); flow rate: 1.0 mL/min; detection: UV 210 nm

Racemic

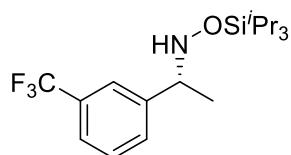


Peak	RT	Area	Area %
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1	11.101	7.674e3	50.245
2	13.514	7.600e3	49.755

Chiral

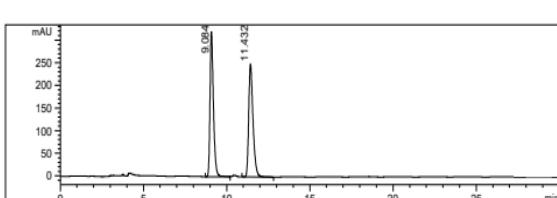


Peak	RT	Area	Area %
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1	11.174	4.954e3	18.785
2	13.467	2.142e4	81.215



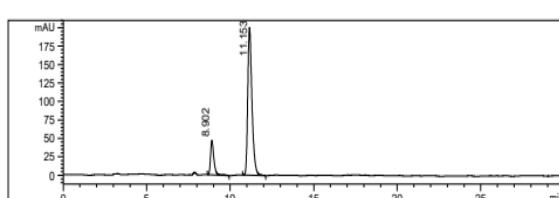
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Racemic

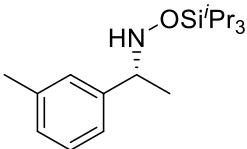


Peak	RT	Area	Area %
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1	9.084	4.563e3	50.263
2	11.432	4.515e3	49.737

Chiral



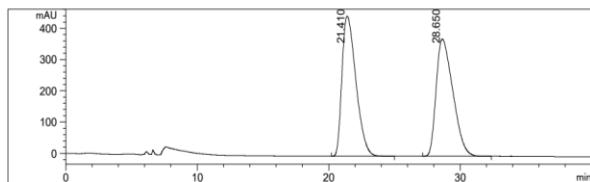
Peak	RT	Area	Area %
#	[min]	-----	-----
1	8.902	658.330	15.843
2	11.153	3.497e3	84.157



2o

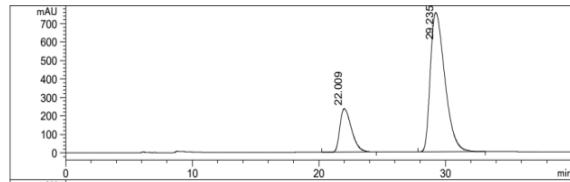
HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (97/3); flow rate: 1.0 mL/min; detection: UV 210 nm

Racemic

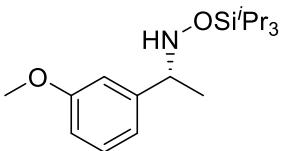


Peak	RT	Area	Area %
#	[min]	-----	-----
1	21.410	3.306e4	50.014
2	28.650	3.304e4	49.986

Chiral



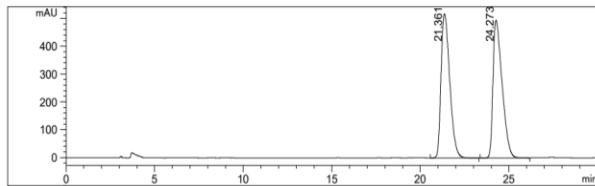
Peak	RT	Area	Area %
#	[min]	-----	-----
1	22.009	1.495e4	19.928
2	29.235	6.006e4	80.072



2p

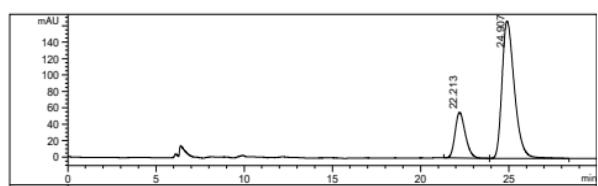
HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (92/8); flow rate: 1.0 mL/min; detection: UV 210 nm

Racemic

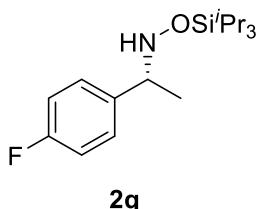


Peak	RT	Area	Area %
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1	21.361	1.768e4	50.014
2	24.273	1.767e4	49.986

Chiral

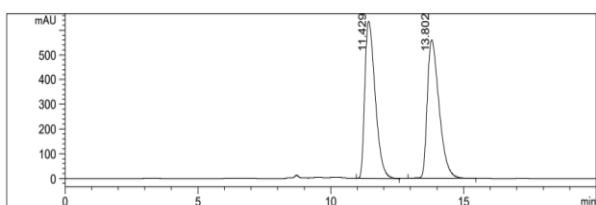


Peak	RT	Area	Area %
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1	22.213	2.253e3	21.579
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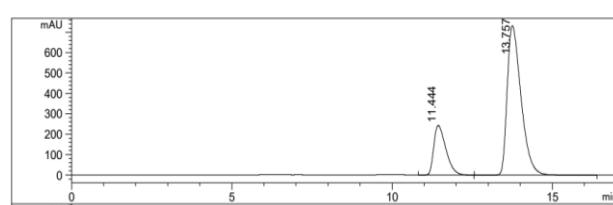


HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (97/3); flow rate: 1.0 mL/min; detection: UV 210 nm

Racemic

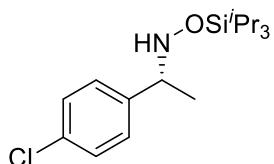


Chiral



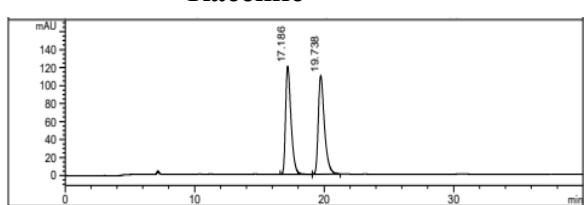
Peak	RT [min]	Area	Area %
1	11.429	1.724e4	49.934
2	13.802	1.729e4	50.066

Peak	RT [min]	Area	Area %
1	11.444	6.382e3	22.453
2	13.757	2.204e4	77.547

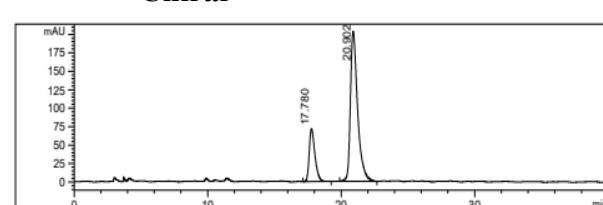


HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (98/2); flow rate: 1.0 mL/min; detection: UV 210 nm

Racemic

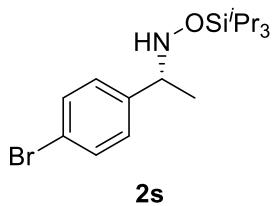


Chiral



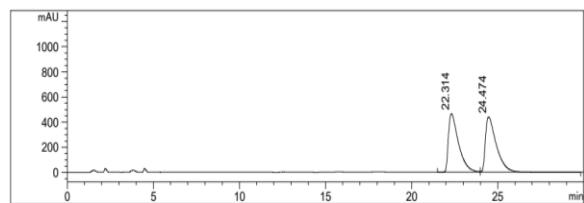
Peak	RT [min]	Area	Area %
1	17.186	3.493e3	49.205
2	19.738	3.606e3	50.795

Peak	RT [min]	Area	Area %
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2	20.902	7.579e3	78.201



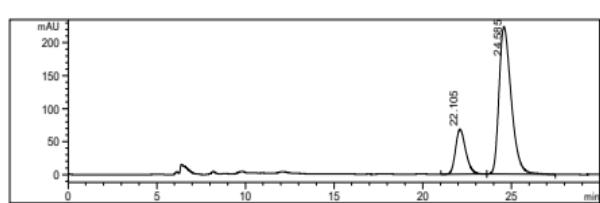
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Racemic

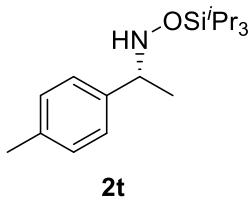


Peak	RT	Area	Area %
#	[min]	-----	-----
1	22.314	1.907e4	49.499
2	24.474	1.946e4	50.501

Chiral

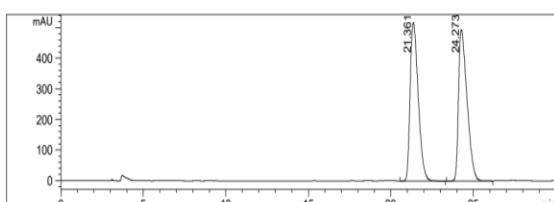


Peak	RT	Area	Area %
#	[min]	-----	-----
1	22.105	2.721e3	20.334
2	24.585	1.066e4	79.666



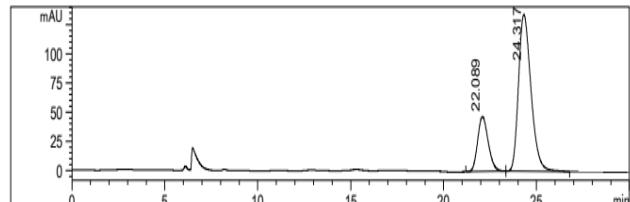
HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (97/3); flow rate: 1.0 mL/min; detection: UV 210 nm

Racemic

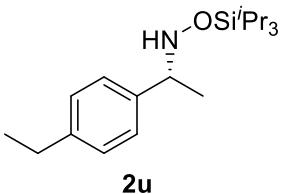


Peak	RT	Area	Area %
#	[min]	-----	-----
1	21.361	1.768e4	50.014
2	24.273	1.767e4	49.986

Chiral

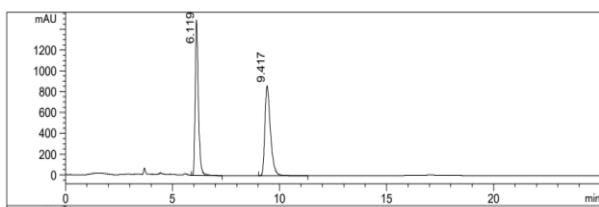


Peak	RT	Area	Area %
#	[min]	-----	-----
1	22.090	1.027e3	22.999
2	24.317	3.437e3	77.001



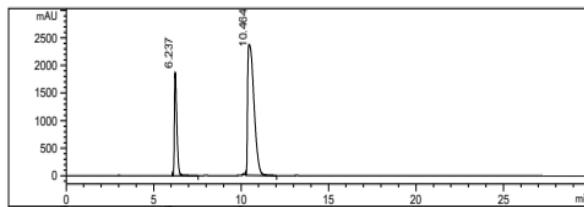
HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (95/5); flow rate: 1.0 mL/min; detection: UV 210 nm

Racemic

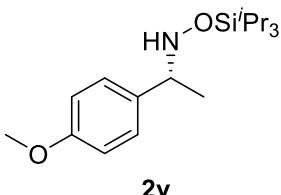


Peak	RT	Area	Area %
#	[min]	-----	-----
1	6.119	1.540e4	49.858
2	9.417	1.549e4	50.142

Chiral

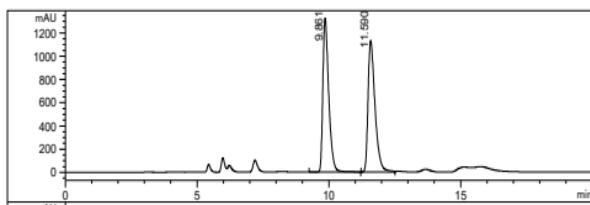


Peak	RT	Area	Area %
#	[min]	-----	-----
1	6.237	1.810e4	23.177
2	10.464	5.999e4	76.823



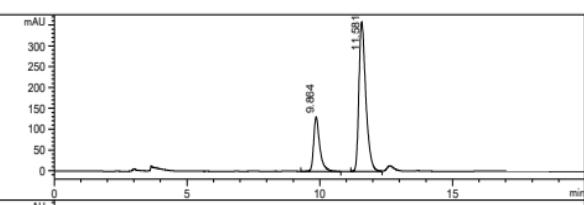
HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (92/8); flow rate: 1.0 mL/min; detection: UV 210 nm

Racemic

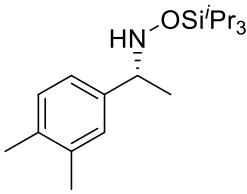


Peak	RT	Area	Area %
#	[min]	-----	-----
1	9.861	2.053e4	49.708
2	11.590	2.077e4	50.292

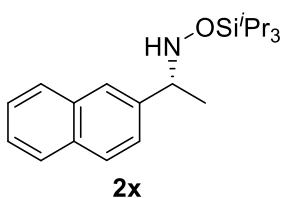
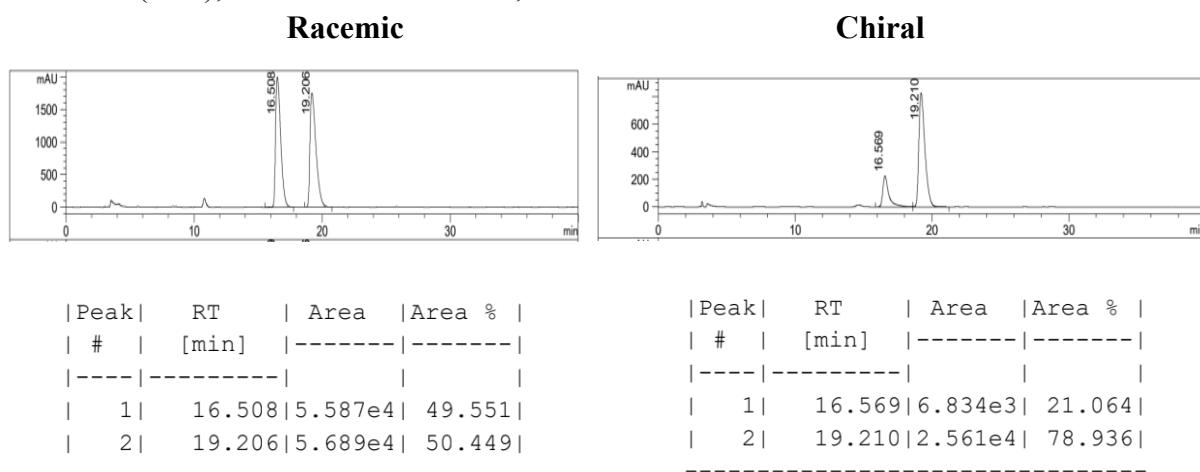
Chiral



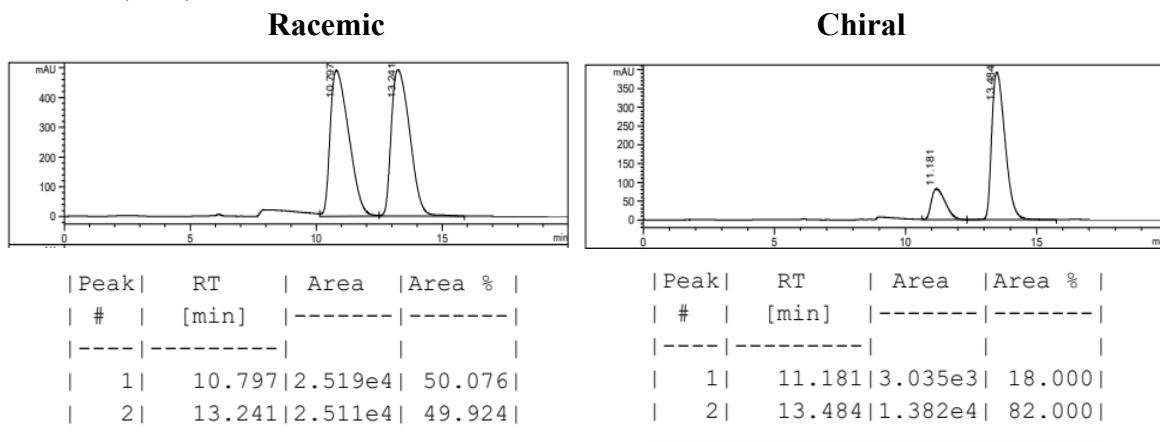
Peak	RT	Area	Area %
#	[min]	-----	-----
1	9.864	2.074e3	24.273
2	11.581	6.471e3	75.727

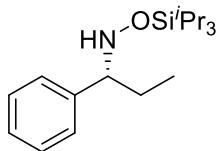


HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (95/5); flow rate: 1.0 mL/min; detection: UV 210 nm



HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (97/3); flow rate: 1.0 mL/min; detection: UV 210 nm

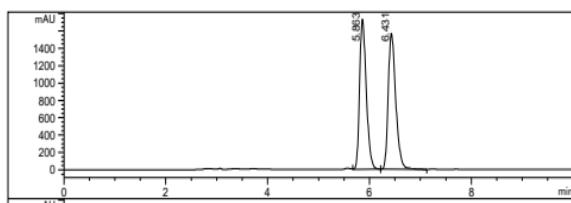




2y

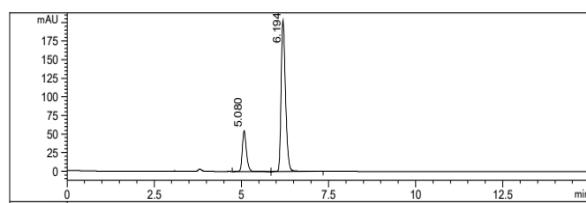
HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (97/3); flow rate: 1.0 mL/min; detection: UV 210 nm

Racemic

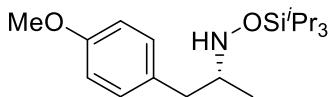


Peak	RT	Area	Area %
#	[min]	-----	-----
1	5.863	1.581e4	49.565
2	6.431	1.609e4	50.435

Chiral



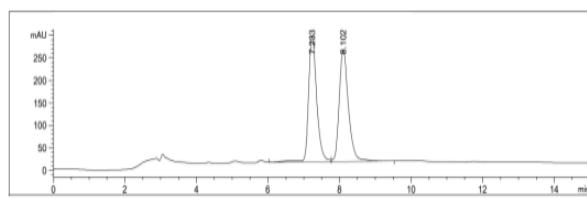
Peak	RT	Area	Area %
#	[min]	-----	-----
1	5.080	445.616	20.061
2	6.194	1.776e3	79.939



2z

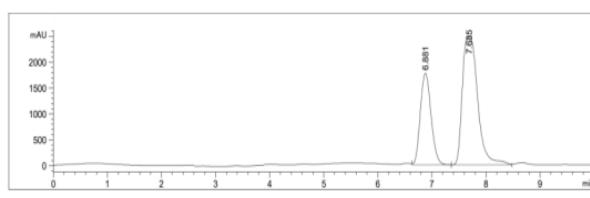
HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (80/20); flow rate: 1.0 mL/min; detection: UV 210 nm

Racemic

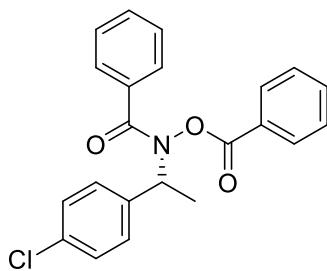


Peak	RT	Area	Area %
#	[min]	-----	-----
1	7.233	1.947e4	50.147
2	8.102	1.887e4	49.852

Chiral

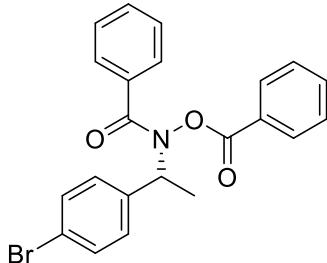
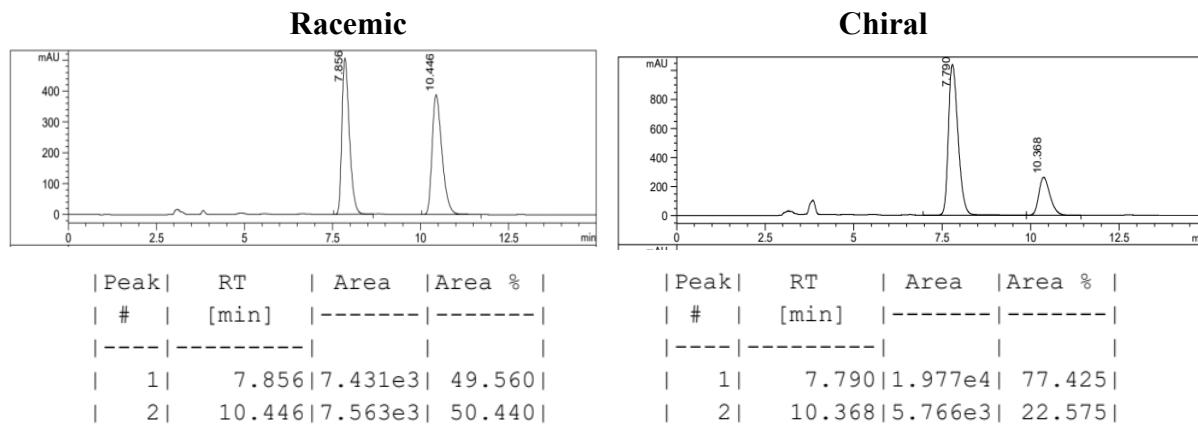


Peak	RT	Area	Area %
#	[min]	-----	-----
1	6.881	2.443e4	33.316
2	7.685	4.890e4	66.684



7

HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (90/10); flow rate: 1.0 mL/min; detection: UV 210 nm



8

HPLC Conditions: Column: Chiralcel OD-H, Daicel Chemical Industries, Ltd., Eluent: Hexanes/IPA (90/10); flow rate: 1.0 mL/min; detection: UV 210 nm

