

Supporting Information

Efficient Access to Chiral 1,2-Amino Alcohols *via* Ir/f-Amphox-Catalyzed Asymmetric Hydrogenation of α -Amino Ketones

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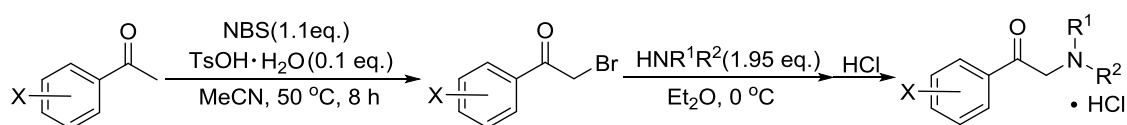
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1. General remarks:

All reactions and manipulations which are sensitive to moisture or air were performed in an argon-filled glovebox or using standard Schlenk techniques. Hydrogen gas (99.999%) was purchased from Shanghai Regulator Factory Co., Ltd. Anhydrous *i*-PrOH and EtOH were purchased from J&K. Anhydrous CF₃CH₂OH were purchased from Sigma-Aldrich, Anhydrous THF, 1,4-dioxane and hexane was distilled from sodium benzophenone ketyl. Anhydrous ClCH₂CH₂Cl were freshly distilled from calcium hydride. Na₂CO₃, K₂CO₃, Cs₂CO₃, NaHCO₃, NaOH, KOH, *t*-BuONa and *t*-BuOK was purchased from J&K. [Ir(COD)Cl]₂ was prepared according to the literature.¹⁻² ¹H, ¹³C and ³¹P NMR spectra were recorded with a Bruker ADVANCE III (400 MHz) spectrometer with CDCl₃ as the solvent and tetramethylsilane (TMS) as the internal standard. Chemical shifts are reported in parts per million (ppm, δ scale) downfield from TMS at 0.00 ppm and referenced to the CDCl₃ at 7.26 ppm (for ¹H NMR) or 77.0 ppm (for ¹³C NMR). Data are reported as: multiplicity (s = singlet, d = doublet, t = triplet, q = quartet, m = multiplet), coupling constant in hertz (Hz) and signal area integration in natural numbers. ¹³C NMR and ³¹P NMR analyses were run with decoupling. Optical rotations [α]_D were determined using a PERKIN ELMER polarimeter 343 instrument. HPLC analyses were performed using Daicel chiral column.

All the α-amino ketones were prepared according the literature.³⁻⁷ The absolute configuration of products were determined by comparison of analytical data with the literature (HPLC spectra, optical rotation).^{6,9-12} The absolute configuration of others were assigned by analogy.

2. General procedure for the preparation of α-amino ketones.

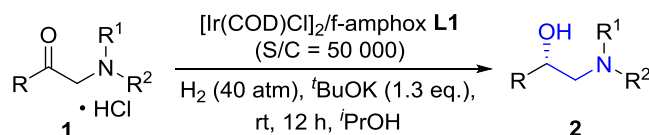


Step 1:³⁻⁴ N-bromosuccinimide was added to the stirred solution of acetophenone in acetonitrile. The resulting reaction mixture was stirred for 10-15 minutes. After that *p*-TsOH·H₂O was added to the reaction mixture and refluxed for 8 h and monitored by TLC. After completion of the reaction, reaction content was brought to room temperature and washed with saturated solution of sodium bicarbonate and extracted with ethyl acetate, organic layer was dried over sodium sulphate and concentrated under reduced pressure. The obtained residues were used for the next step directly.

Step 2:⁵⁻⁷ The obtained residues of step 1 was added to the stirred solution of NHR¹R² in diethylether, then stirred for 3-5 h and monitored by TLC. After completion of the reaction, the mixture was filtered and removed the solvent under reduced pressure. The residue was washed with saturated solution of sodium bicarbonate and the aqueous phase was extracted with ethyl

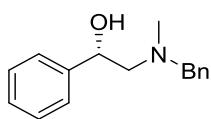
acetate. The combined organic extractions were dried over Na₂SO₄ and concentrated under reduced pressure to yield a residue. The residue was purified by flash chromatography on silica gel with ethyl acetate to get the corresponding α -amino ketones. The obtained α -amino ketone was then treated with hydrochloric acid and recrystallized in hot anhydrous EtOH to give the corresponding hydrochloride as a white crystalline solid.

3. Asymmetric hydrogenation of α -amino ketones:^{3,6,8}



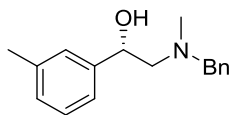
General procedure (at S/C = 50 000): To a 4.0 mL vial was added the catalyst precursor [Ir(COD)Cl]₂ (1.4 mg, 2.0×10⁻³ mmol), ligand **L1** (2.4 mg, 4.2×10⁻³ mmol) and anhydrous *i*-PrOH (2.0 mL) in the argon-filled glovebox. The mixture was stirred for 2.0 h at 25 °C giving orange red solution. And then 0.2 mmol of α -amino ketone (as a hydrochloride), *t*-BuOK (29.2 mg, 1.3 eq.) were added into a 5 mL hydrogenation vessel. 1.0 mL anhydrous *i*-PrOH was added as solvent and a solution of Ir/f-amphox **L1** in anhydrous *i*-PrOH (2.0 μ L) was added *via* an injection port. Then the vessel was placed in an autoclave, closed it and moved it out from glovebox. The autoclave quickly purged with hydrogen gas for three times, then pressurized to 40 atm H₂. The reaction solution was stirred at room temperature (25 °C-30 °C) until for 12 h, then released pressure carefully. The solution of reaction mixture was purified by a flash chromatography on a silical gel with ethyl acetate and the solvent was removed under reduced pressure. The ee value was determined by chiral HPLC analysis of the chiral amino alcohol directly. The characterization data of compounds **2a-2n** are in accordance with the reported data in the literature.^{6,9-12}

(*S*)-2-(Benzyl(methyl)amino)-1-phenylethanol (**2a**)



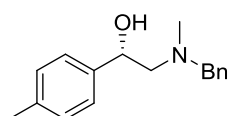
Light yellow oil; >99% yield; >99% ee. The enantiomeric excess was determined by HPLC on Chiralcel OJ-H column, 210 nm, 25 °C, *n*-hexane: *i*-PrOH = 95:5; flow 1.0 mL/min; *t_R* (major) = 14.3 min, *t_R* (minor) = 16.8 min. [α]_D²⁵ = 50.6 (*c* 1.0, EtOH). ¹H NMR (400 MHz, CDCl₃) δ 7.28-7.16 (m, 10H), 4.67 (dd, *J* = 10.4, 3.6 Hz, 1H), 3.67 (d, *J* = 13.1 Hz, 1H), 3.46 (d, *J* = 13.1 Hz, 1H), 2.53-2.43 (m, 2H), 2.24 (s, 3H). ¹³C NMR (101 MHz, CDCl₃) δ 142.16, 138.17, 129.11, 128.46, 127.52, 127.39, 125.94, 69.39, 65.50, 62.38, 41.75.

(*S*)-2-(Benzyl(methyl)amino)-1-(3-methylphenyl)ethanol (**2b**)



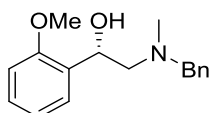
Light yellow oil; >99% yield; >99% ee. The enantiomeric excess was determined by HPLC on Chiracel OJ-H column, 210 nm, 25 °C, *n*-hexane: *i*-PrOH = 95:5; flow 1.0 mL/min; t_R (major) = 11.3 min, t_R (minor) = 12.8 min. $[\alpha]_D^{25} = 22.4$ (*c* 0.5, EtOH). $^1\text{H NMR}$ (400 MHz, CDCl_3) δ 7.43-7.24 (m, 5H), 7.23-7.03 (m, 4H), 4.72 (dd, $J = 10.5, 3.4$ Hz, 1H), 3.74 (d, $J = 13.1$ Hz, 1H), 3.53 (d, $J = 13.1$ Hz, 1H), 2.58-2.50 (m, 1H), 2.34 (s, 3H), 2.31 (s, 3H). $^{13}\text{C NMR}$ (101 MHz, CDCl_3) δ 142.08, 138.02, 129.12, 127.38, 126.58, 123.03, 69.39, 65.56, 62.39, 41.71, 21.51.

(S)-2-(Benzyl(methyl)amino)-1-(4-methylphenyl)ethanol (2c)



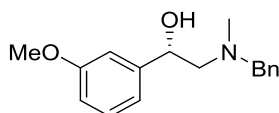
Light yellow oil; >99% yield; >99% ee. The enantiomeric excess was determined by HPLC on Chiracel OJ-H column, 210 nm, 25 °C, *n*-hexane: *i*-PrOH = 85:15; flow 1.0 mL/min; t_R (major) = 7.7 min, t_R (minor) = 8.7 min. $[\alpha]_D^{25} = 63.25$ (*c* 1.2, EtOH). $^1\text{H NMR}$ (400 MHz, CDCl_3) δ 7.32-7.23 (m, 7H), 7.14 (d, $J = 7.9$ Hz, 2H), 4.72 (dd, $J = 10.5, 3.4$ Hz, 1H), 3.74 (d, $J = 13.1$ Hz, 1H), 3.52 (d, $J = 13.1$ Hz, 1H), 2.51-2.43 (m, 2H), 2.33 (s, 3H), 2.31 (s, 3H). $^{13}\text{C NMR}$ (101 MHz, CDCl_3) δ 139.12, 138.18, 137.15, 129.11, 129.06, 128.44, 127.37, 125.90, 69.24, 65.56, 62.38, 41.73, 21.19.

(S)-2-(Benzyl(methyl)amino)-1-(2-methoxyphenyl)ethanol (2d)



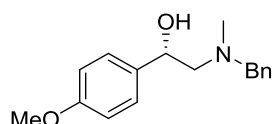
Light yellow oil; >99% yield; >99% ee. The enantiomeric excess was determined by HPLC on Chiracel OJ-H column, 210 nm, 25 °C, *n*-hexane: *i*-PrOH = 95:5; flow 1.0 mL/min; t_R (major) = 13.3 min, t_R (minor) = 15.3 min. $[\alpha]_D^{25} = 84.08$ (*c* 1.2, EtOH). $^1\text{H NMR}$ (400 MHz, CDCl_3) δ 7.52 (dd, $J = 7.5, 1.3$ Hz, 1H), 7.42-7.18 (m, 6H), 6.97-6.95 (m, 1H), 6.83 (d, $J = 8.2$ Hz, 1H), 5.16 (dd, $J = 10.0, 3.1$ Hz, 3H), 3.87 (brs, 1H), 3.79 (s, 3H), 3.75 (d, $J = 13.1$ Hz, 1H), 3.53 (d, $J = 13.1$ Hz, 1H), 2.69 (dd, $J = 12.3, 3.2$ Hz, 1H), 2.50-2.47 (m, 1H), 2.32 (s, 3H). $^{13}\text{C NMR}$ (101 MHz, CDCl_3) δ 156.25, 138.21, 130.39, 129.22, 128.38, 128.11, 127.29, 126.51, 120.81, 110.07, 64.49, 63.44, 62.16, 55.30, 41.97.

(S)-2-(Benzyl(methyl)amino)-1-(3-methoxyphenyl)ethanol (2e)



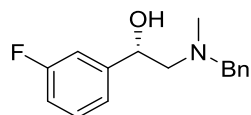
Light yellow oil; >99% yield; >99% ee. The enantiomeric excess was determined by HPLC on Chiracel OJ-H column, 210 nm, 25 °C, *n*-hexane: *i*-PrOH = 85:15; flow 1.0 mL/min; t_R (major) = 10.9 min, t_R (minor) = 11.5 min. $[\alpha]_D^{25} = 44.40$ (*c* 0.5, EtOH). $^1\text{H NMR}$ (400 MHz, CDCl_3) δ 7.39-7.32 (m, 3H), 7.31-7.19 (m, 3H), 6.93-6.90 (m, 2H), 6.82-6.79 (m, 1H), 4.73 (dd, $J = 10.2, 3.7$ Hz, 1H), 3.80 (s, 3H), 3.74 (d, $J = 13.1$ Hz, 1H), 3.53 (d, $J = 13.1$ Hz, 1H), 2.60-2.51 (m, 2H), 2.31 (s, 3H). $^{13}\text{C NMR}$ (101 MHz, CDCl_3) δ 159.73, 143.94, 138.15, 129.36, 129.11, 128.45, 127.38, 118.22, 113.13, 111.17, 69.27, 65.41, 62.36, 55.24, 41.74.

(S)-2-(Benzyl(methyl)amino)-1-(4-methoxyphenyl)ethanol (2f)



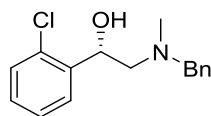
Light yellow oil; 99% yield; >99% ee. The enantiomeric excess was determined by HPLC on Chiracel OJ-H column, 220 nm, 25 °C, *n*-hexane: *i*-PrOH = 85:15; flow 1.0 mL/min; t_R (major) = 14.6 min, t_R (minor) = 17.7 min. $[\alpha]_D^{25} = 46.40$ (*c* 1.0, EtOH). $^1\text{H NMR}$ (400 MHz, CDCl_3) δ 7.36-7.28 (m, 7H), 6.92-6.85 (m, 2H), 4.70 (dd, $J = 10.6, 3.3$ Hz, 1H), 3.79 (d, $J = 5.6$ Hz, 3H), 3.81-3.72 (m, 1H), 3.53 (d, $J = 13.1$ Hz, 1H), 2.62-2.56 (m, 1H), 2.48 (dd, $J = 12.4, 3.4$ Hz, 1H), 2.31 (s, 3H). $^{13}\text{C NMR}$ (101 MHz, CDCl_3) δ 159.04, 138.22, 134.16, 129.09, 128.43, 127.35, 127.20, 113.76, 69.02, 65.52, 62.38, 55.30, 41.74.

(S)-2-(benzyl(methyl)amino)-1-(3-fluorophenyl)ethanol (2g)



Light yellow oil; >99% yield; >99% ee. The enantiomeric excess was determined by HPLC on Chiracel OJ-H column, 210 nm, 25 °C, *n*-hexane: *i*-PrOH = 97:3; flow 1.0 mL/min; t_R (major) = 15.2 min, t_R (minor) = 16.2 min. $[\alpha]_D^{25} = 48.80$ (*c* 1.0, EtOH). $^1\text{H NMR}$ (400 MHz, CDCl_3) δ 7.45-7.21 (m, 6H), 7.10-7.08 (m, 2H), 6.98-6.88 (m, 1H), 4.73 (dd, $J = 8.7, 5.3$ Hz, 1H), 3.74 (d, $J = 13.0$ Hz, 1H), 3.53 (d, $J = 13.1$ Hz, 1H), 2.56-2.52 (m, 2H), 2.32 (s, 3H). $^{13}\text{C NMR}$ (101 MHz, CDCl_3) δ 162.99 (d, $J = 246.4$ Hz), 144.99 (d, $J = 7.1$ Hz), 137.99, 129.08, 128.48, 127.46, 121.41 (d, $J = 7.1$ Hz), 114.28 (d, $J = 22.22$ Hz), 112.81 (d, $J = 22.22$ Hz), 68.80, 65.20, 62.34, 41.75.

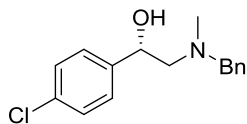
(S)-2-(Benzyl(methyl)amino)-1-(2-chlorophenyl)ethanol (2h)



Light yellow oil; >99% yield; >99% ee. The enantiomeric excess was determined by HPLC on Chiracel AD-H column, 210 nm, 25 °C, *n*-hexane: *i*-PrOH = 98:2; flow 0.3 mL/min; t_R (minor) = 32.9 min, t_R (major) = 36.0 min. $[\alpha]_D^{25} = 100.1$ (*c* 1.0, EtOH). $^1\text{H NMR}$ (400 MHz, CDCl_3) δ

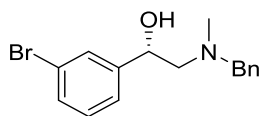
7.54 (dd, $J = 7.7, 1.3$ Hz, 1H), 7.34-7.07 (m, 8H), 5.10 (dd, $J = 10.3, 2.9$ Hz, 1H), 3.92 (brs, 1H), 3.69 (d, $J = 13.1$ Hz, 1H), 3.44 (d, $J = 13.1$ Hz, 1H), 2.68-2.65 (m, 1H), 2.35-2.32 (m, 1H), 2.26 (s, 3H). ^{13}C NMR (101 MHz, CDCl_3) δ 139.57, 138.16, 131.72, 129.20, 129.13, 128.47, 128.42, 127.40, 127.34, 127.18, 66.49, 63.22, 62.25, 41.85.

(S)-2-(Benzyl(methyl)amino)-1-(4-chlorophenyl)ethanol (2i)



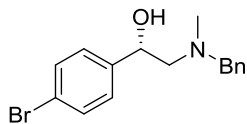
White solid; >99% yield; >99% ee. The enantiomeric excess was determined by HPLC on Chiralcel OJ-H column, 220 nm, 25 °C, *n*-hexane: *i*-PrOH = 85:15; flow 1.0 mL/min; t_{R} (major) = 7.8 min, t_{R} (minor) = 8.3 min. $[\alpha]_{\text{D}}^{25} = 43.00$ (*c* 1.0, EtOH). ^1H NMR (400 MHz, CDCl_3) δ 7.38-7.26 (m, 9H), 4.71 (dd, $J = 9.6, 4.5$ Hz, 1H), 3.74 (d, $J = 13.0$ Hz, 1H), 3.53 (d, $J = 13.0$ Hz, 1H), 2.56-2.48 (m, 2H), 2.32 (s, 3H). ^{13}C NMR (101 MHz, CDCl_3) δ 140.69, 138.01, 133.09, 129.08, 128.50, 128.48, 127.44, 127.27, 68.76, 65.29, 62.35, 41.75.

(S)-2-(Benzyl(methyl)amino)-1-(3-bromophenyl)ethanol (2j)



Colorless oil; >99% yield; >99% ee. The enantiomeric excess was determined by HPLC on Chiralcel OJ-H column, 210 nm, 25 °C, *n*-hexane: *i*-PrOH = 97:3; flow 1.0 mL/min; t_{R} (minor) = 18.2 min, t_{R} (major) = 20.2 min. $[\alpha]_{\text{D}}^{25} = 29.41$ (*c* 1.0, EtOH). ^1H NMR (400 MHz, CDCl_3) δ 7.52 (s, 1H), 7.42-7.23 (m, 7H), 7.19 (t, $J = 7.7$ Hz, 1H), 4.70 (dd, $J = 8.6, 5.4$ Hz, 1H), 3.73 (d, $J = 13.1$ Hz, 1H), 3.53 (d, $J = 13.1$ Hz, 1H), 2.60-2.49 (m, 2H), 2.31 (s, 3H). ^{13}C NMR (101 MHz, CDCl_3) δ 144.63, 137.98, 130.54, 129.95, 129.08, 128.97, 128.49, 127.47, 124.50, 122.54, 68.76, 65.27, 62.35, 41.72.

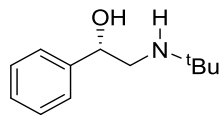
(S)-2-(Benzyl(methyl)amino)-1-(4-bromophenyl)ethanol (2k)



White solid; >99% yield; >99% ee. The enantiomeric excess was determined by HPLC on Chiralcel OJ-H column, 210 nm, 25 °C, *n*-hexane: *i*-PrOH = 95:5; flow 1.0 mL/min; t_{R} (major) = 16.1 min, t_{R} (minor) = 17.4 min. $[\alpha]_{\text{D}}^{25} = 38.40$ (*c* 1.0, EtOH). ^1H NMR (400 MHz, CDCl_3) δ 7.37 (d, $J = 8.2$ Hz, 2H), 7.32-7.18 (m, 5H), 7.14 (d, $J = 8.1$ Hz, 2H), 4.62 (dd, $J = 9.1, 4.5$ Hz, 1H), 3.66 (d, $J = 13.0$ Hz, 1H), 3.45 (d, $J = 13.0$ Hz, 1H), 2.45-2.41 (m, 2H), 2.24 (s, 3H). ^{13}C NMR

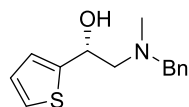
(101 MHz, CDCl₃) δ 140.18, 136.91, 130.37, 127.41, 126.56, 126.38, 120.13, 67.73, 64.15, 61.27, 40.68.

(S)-2-(tert-butylamino)-1-phenylethanol (2l)



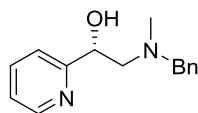
White solid; >99% yield; >99% ee. The enantiomeric excess was determined by HPLC on Chiracel AD-H column, 210 nm, 25 °C, *n*-hexane: *i*-PrOH = 85:15; flow 1.0 mL/min; t_R (major) = 12.5 min, t_R (minor) = 15.6 min. HPLC conditions (To the corresponding benzoyl derivatives). $[\alpha]_D^{25} = 25.9$ (*c* 1.2, EtOH). ¹H NMR (400 MHz, CDCl₃) δ 7.51-7.45 (m, 2H), 7.40-7.28 (m, 3H), 5.45 (dd, *J* = 10.6, 2.0 Hz, 1H), 3.24 (dd, *J* = 12.2, 2.3 Hz, 1H), 3.06 (dd, *J* = 12.0, 10.8 Hz, 1H), 1.51 (s, 9H). ¹³C NMR (101 MHz, CDCl₃) δ 139.87, 128.74, 128.29, 125.85, 68.74, 58.10, 49.66, 26.07.

(R)-2-(benzyl(methyl)amino)-1-(thiophen-2-yl)ethanol (2m)



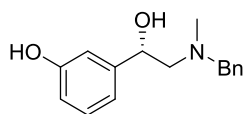
Colorless oil; >99% yield; >99% ee. The enantiomeric excess was determined by HPLC on Chiracel OJ-H column, 210 nm, 25 °C, *n*-hexane: *i*-PrOH = 95:5; flow 1.0 mL/min; t_R (minor) = 19.2 min, t_R (major) = 21.6 min. $[\alpha]_D^{25} = 14.7$ (*c* 1.2, EtOH). ¹H NMR (400 MHz, CDCl₃) δ 7.40-7.21 (m, 6H), 7.01-6.93 (m, 2H), 5.02 (dd, *J* = 10.4, 3.5 Hz, 1H), 3.73 (d, *J* = 13.0 Hz, 1H), 3.56 (d, *J* = 13.1 Hz, 1H), 2.77 (dd, *J* = 12.3, 10.4 Hz, 1H), 2.64 (dd, *J* = 12.4, 3.5 Hz, 1H), 2.30 (s, 3H). ¹³C NMR (101 MHz, CDCl₃) δ 145.78, 138.04, 129.09, 128.47, 127.42, 126.62, 124.46, 123.73, 65.94, 65.36, 62.38, 41.72.

(R)-2-(benzyl(methyl)amino)-1-(pyridin-2-yl)ethanol (2n)



Light brown oil; 99% yield; >99% ee. The enantiomeric excess was determined by HPLC on Chiracel AS-H column, 210 nm, 25 °C, *n*-hexane: *i*-PrOH = 97:3; flow 1.0 mL/min; t_R (minor) = 12.9 min, t_R (major) = 17.3 min. $[\alpha]_D^{25} = 80.8$ (*c* 1.2, EtOH). ¹H NMR (400 MHz, CDCl₃) δ 8.52 (d, *J* = 4.4 Hz, 1H), 7.68 (td, *J* = 7.7, 1.7 Hz, 1H), 7.51 (d, *J* = 7.9 Hz, 1H), 7.37-7.22 (m, 5H), 7.17 (dd, *J* = 6.9, 5.4 Hz, 1H), 4.87 (dd, *J* = 9.8, 3.9 Hz, 1H), 3.73 (d, *J* = 13.1 Hz, 1H), 3.54 (d, *J* = 13.1 Hz, 1H), 2.81 (dd, *J* = 12.4, 3.9 Hz, 1H), 2.66 (dd, *J* = 12.4, 9.8 Hz, 1H), 2.32 (s, 3H). ¹³C NMR (101 MHz, CDCl₃) δ 161.72, 148.64, 138.35, 136.77, 129.06, 128.39, 127.28, 122.34, 120.42, 70.20, 63.83, 62.40, 41.87.

(S)-3-(2-(benzyl(methyl)amino)-1-hydroxyethyl)phenol (**2o**)



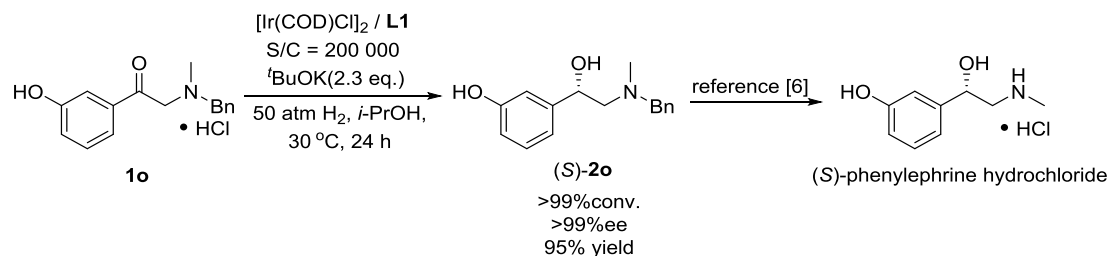
To a 4.0 mL vial was added the catalyst precursor $[\text{Ir}(\text{COD})\text{Cl}]_2$ (1.4 mg, 2.0×10^{-3} mmol), ligand **L1** (2.4 mg, 4.2×10^{-3} mmol) and anhydrous *i*-PrOH (2.0 mL) under argon atmosphere. The mixture was stirred for 2.0 h at 25 °C giving orange red solution in the argon-filled glovebox. And then 0.2 mmol of α -amino ketone **1o** (as a hydrochloride), *t*-BuOK (51.6 mg, 2.3 eq.) were added into a 5 mL hydrogenation vessel. 1 mL anhydrous *i*-PrOH was added as solvent and a solution of Ir/*f*-amphox **L1** in anhydrous *i*-PrOH (2 μL , $c = 2.0 \times 10^{-3}$ mol/L) was added *via* an injection port. Then the vessel was placed in an autoclave, closed it, and moved out of the glovebox. The autoclave was quickly purged with hydrogen gas for three times, then pressurized to 40 atm H_2 . The reaction solution was stirred at room temperature (25 °C - 30 °C) until for 12 h, then released pressure carefully. The reaction solution was treated with 2N HCl aqueous solution to adjust pH to ~7. the solvent was removed under reduced pressure and the residue was purified by a flash chromatography on a silical gel with ethyl acetate (1% Et_3N) as eluent to afford **2o** as a white solid; >99% conv.; 97% yield; >99% ee. The enantiomeric excess was determined by HPLC on Chiracel OJ-H column, 210 nm, 25 °C, *n*-hexane: *i*-PrOH = 80:20; flow 1.0 mL/min; t_{R} (major) = 10.8 min, t_{R} (minor) = 15.9 min. $[\alpha]_{\text{D}}^{25} = 50.7$ (c 1.0, EtOH). ^1H NMR (400 MHz, CDCl_3) δ 7.40-7.24 (m, 5H), 7.16 (t, $J = 7.8$ Hz, 1H), 6.91 (s, 1H), 6.83 (d, $J = 7.6$ Hz, 1H), 6.77-6.68 (m, 1H), 4.74 (dd, $J = 10.3, 3.6$ Hz, 1H), 4.10 (brs, 2H), 3.77 (d, $J = 13.1$ Hz, 1H), 3.57 (d, $J = 13.1$ Hz, 1H), 2.59 (dt, $J = 12.5, 5.4$ Hz, 2H), 2.33 (s, 3H). ^{13}C NMR (101 MHz, CDCl_3) δ 156.17, 143.60, 137.47, 129.57, 129.25, 128.51, 127.56, 117.93, 114.69, 112.83, 69.16, 65.03, 62.18, 41.62.

Asymmetric hydrogenation of α -amino ketone **1a** at S/C = 500 000:

To a 4.0 mL vial was added the catalyst precursor $[\text{Ir}(\text{COD})\text{Cl}]_2$ (1.4 mg, 2.0×10^{-3} mmol), ligand **L1** (2.4 mg, 4.2×10^{-3} mmol) and anhydrous *i*-PrOH (2.0 mL) in the argon-filled glovebox. The mixture was stirred for 2.0 h at 25 °C giving orange red solution. The 10 mmol of α -amino ketone **1a** (as a hydrochloride) and *t*-BuOK (13 mmol, 1.46 g) were added into a 10 ml hydrogenation vessel. 5 mL anhydrous *i*-PrOH was added as solvent and a solution of Ir/*f*-amphox **L1** in anhydrous *i*-PrOH (10 μL , 2.0×10^{-3} mol/L) was added via an injection port. Then the vessel was placed in an autoclave, closed it and moved it out from glovebox. The autoclave was quickly purged with hydrogen gas for three times, then pressurized to 50 atm H_2 . The reaction solution was stirred at 30 °C until no obvious hydrogen pressure drop was observed (48 h), then released pressure carefully. The solvent of the reaction mixture was removed under reduced pressure and the residue was purified by a flash chromatography on a silical gel with ethyl acetate as eluent to

afford the chiral amino alcohol (*S*)-**2a** 2.38 g as a light yellow oil, >99% conversion, >99% yield, >99% ee (*S*).

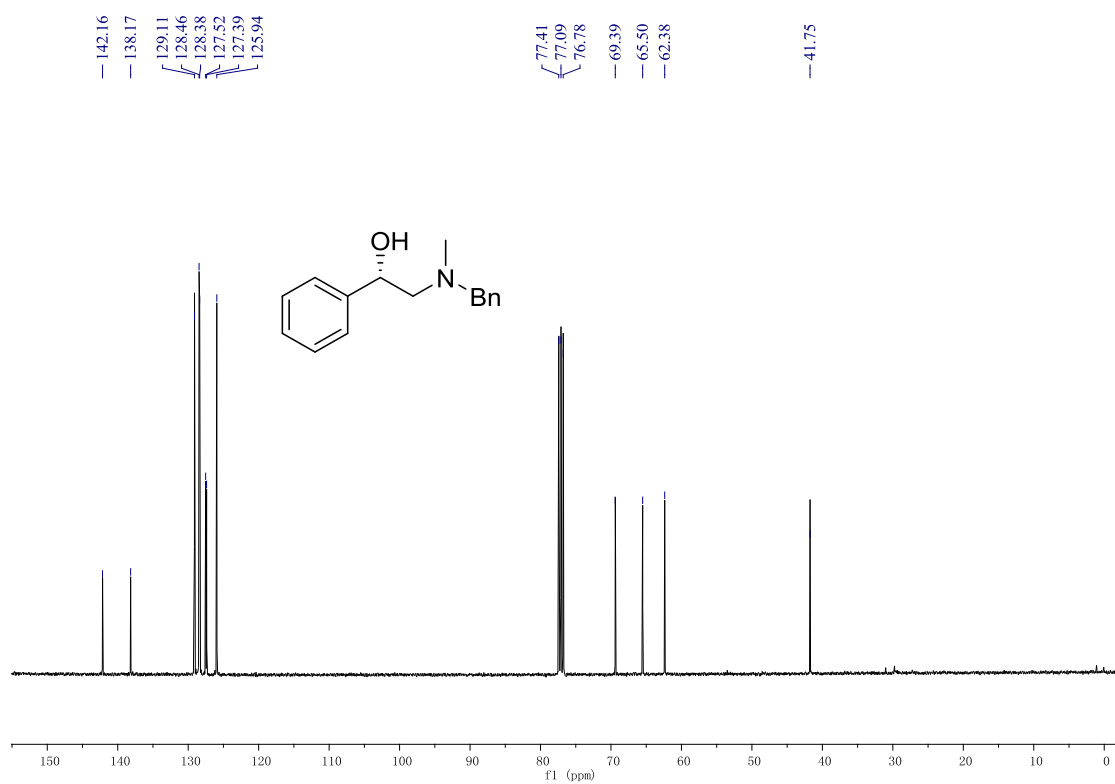
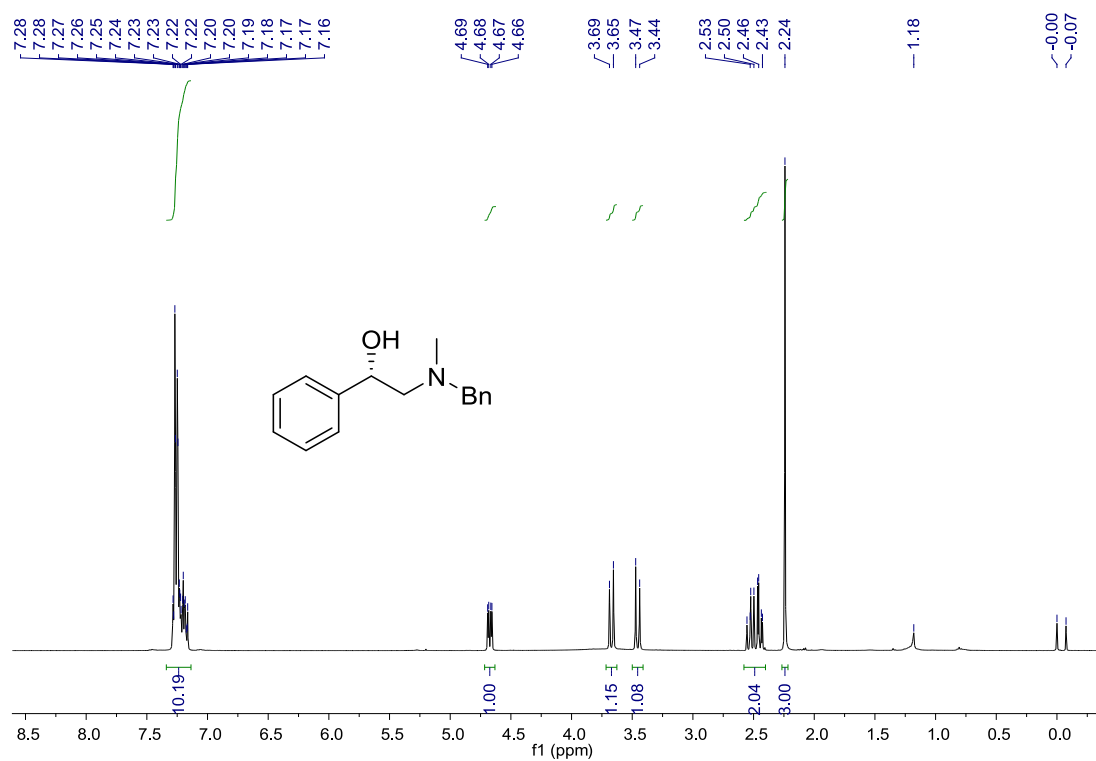
4. The Synthesis of (*S*)-Phenylephrine Hydrochloride at S/C=200 000.^{6,8,13}



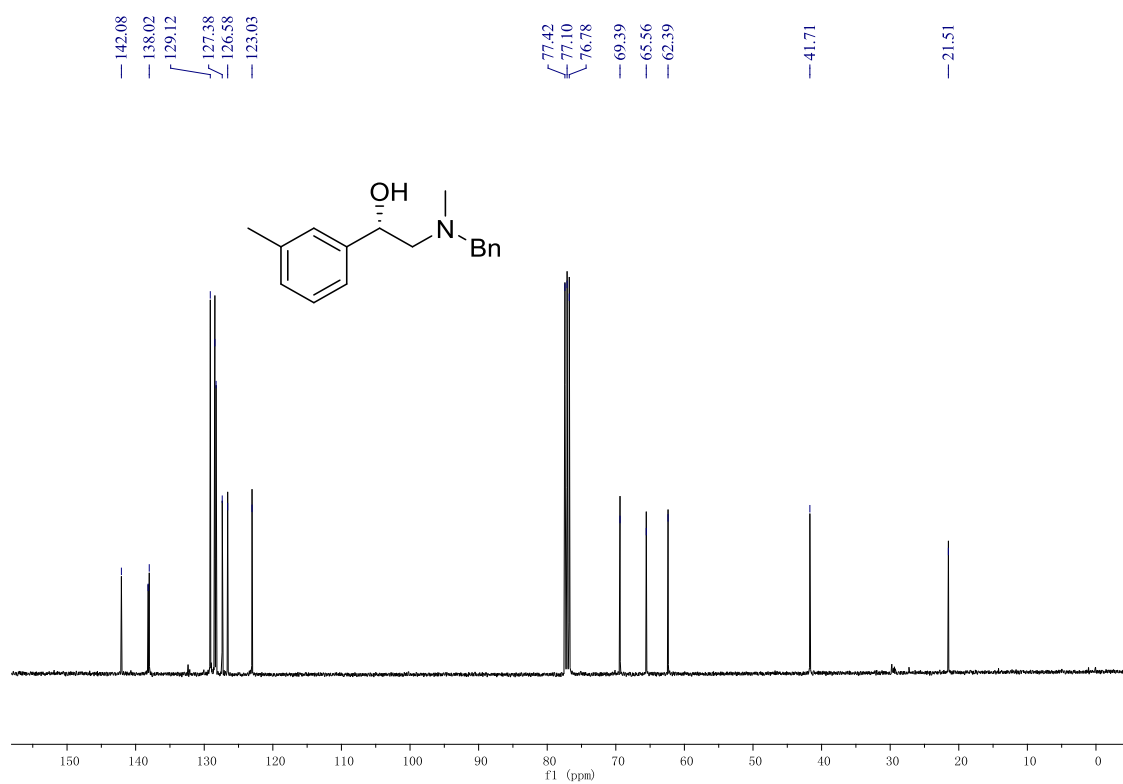
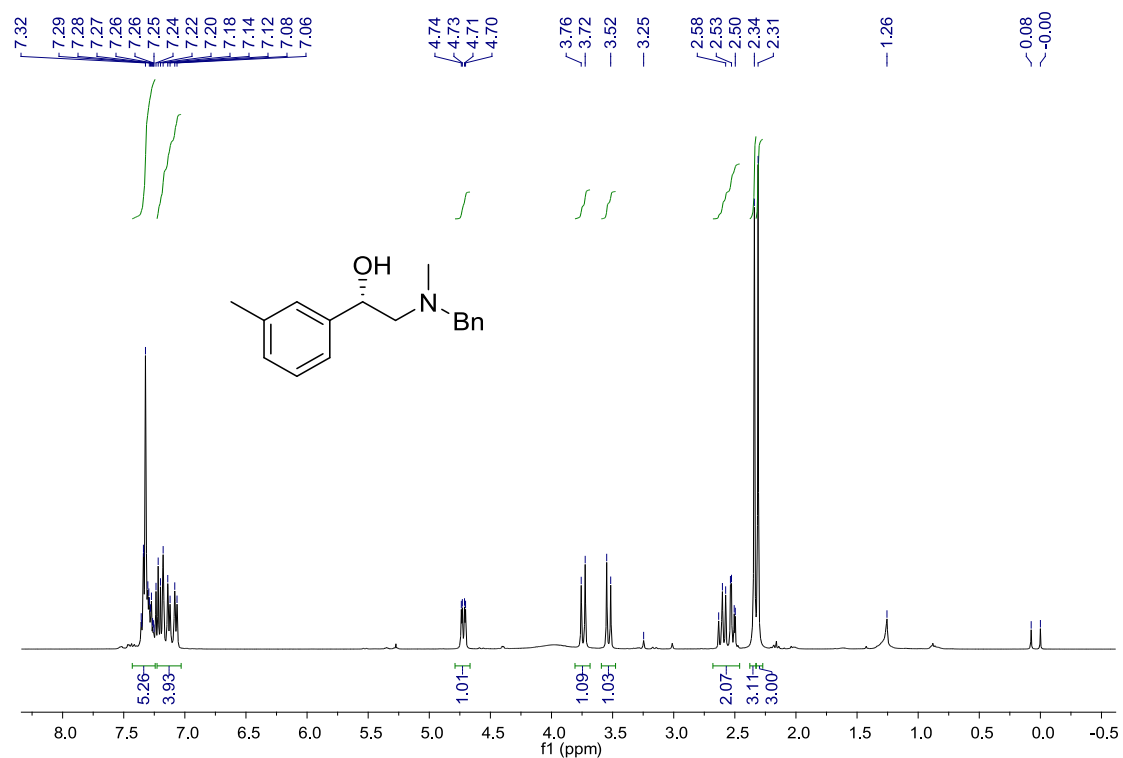
1.17 g (4 mmol) of amino ketone **1o** (as a hydrochloride) and 1.03 g (9.2 mmol) of *t*-BuOK were added into a 10 ml hydrogenation vessel. Then 4 mL of anhydrous *i*-PrOH and 10 μL (2.0×10^{-3} mol/L, 0.02 μmol) of the solution of Ir-L1 in anhydrous *i*-PrOH was added via an injection port. Then the vessel was placed in an autoclave, closed it and moved it out from golvebox. The autoclave was quickly purged with hydrogen gas for three times, then pressurized to 50 atm H_2 . The reaction solution was stirred at 30 °C until no obvious hydrogen pressure drop was observed (24 h), then released pressure carefully. The reaction solution was treated with 2N HCl aqueous solution to adjust pH to ~7. The solvent was removed under reduced pressure and the residue was purified by a flash chromatography on a silical gel with ethyl acetate (1% Et_3N) as eluent to afford (*S*)-**2o** 0.98 g as a white solid; >99% conv.; 95% yield; >99% ee. The enantiomeric excess was determined by HPLC on Chiracel OJ-H column, 210 nm, 25 °C, *n*-hexane: *i*-PrOH = 80:20; flow 1.0 mL/min; t_{R} (major) = 10.8 min, t_{R} (minor) = 15.9 min. $[\alpha]_{\text{D}}^{25} = 50.7$ (*c* 1.0, EtOH). ^1H NMR (400 MHz, CDCl_3) δ 7.40-7.24 (m, 5H), 7.16 (t, $J = 7.8$ Hz, 1H), 6.91 (s, 1H), 6.83 (d, $J = 7.6$ Hz, 1H), 6.77-6.68 (m, 1H), 4.74 (dd, $J = 10.3, 3.6$ Hz, 1H), 4.10 (brs, 2H), 3.77 (d, $J = 13.1$ Hz, 1H), 3.57 (d, $J = 13.1$ Hz, 1H), 2.59 (dt, $J = 12.5, 5.4$ Hz, 2H), 2.33 (s, 3H). ^{13}C NMR (101 MHz, CDCl_3) δ 156.17, 143.60, 137.47, 129.57, 129.25, 128.51, 127.56, 117.93, 114.69, 112.83, 69.16, 65.03, 62.18, 41.62.

5. NMR spectra

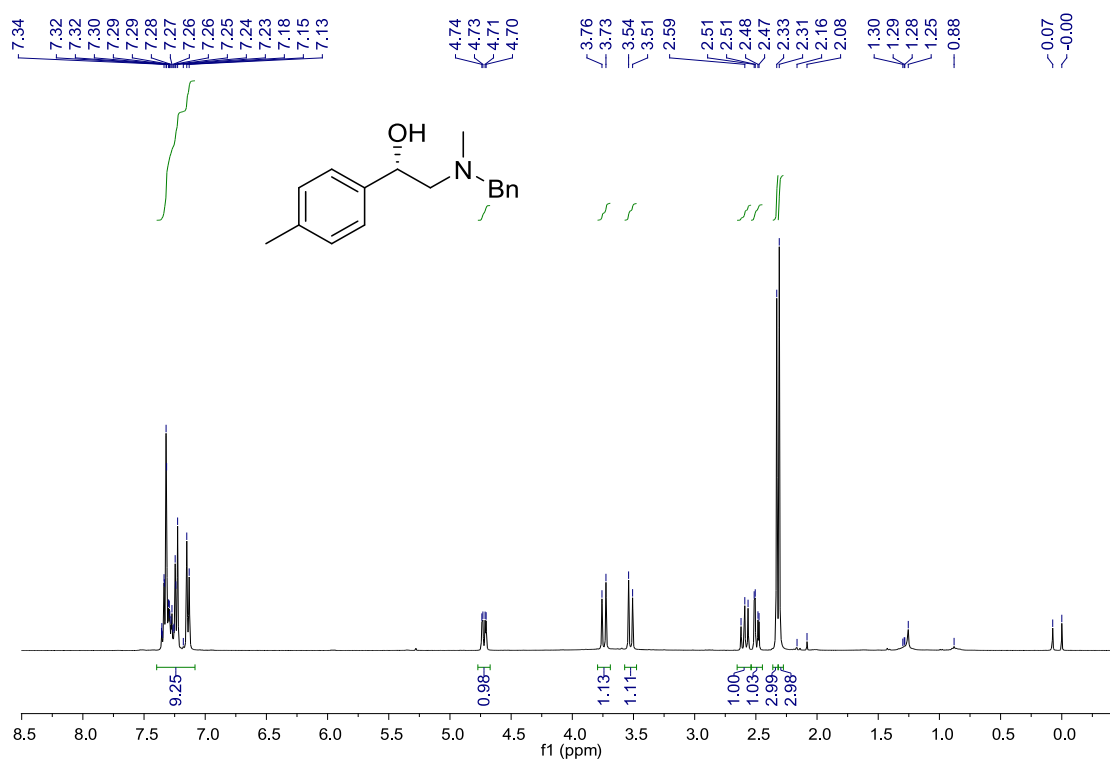
(S)-2-(Benzyl(methyl)amino)-1-phenylethanol (2a)



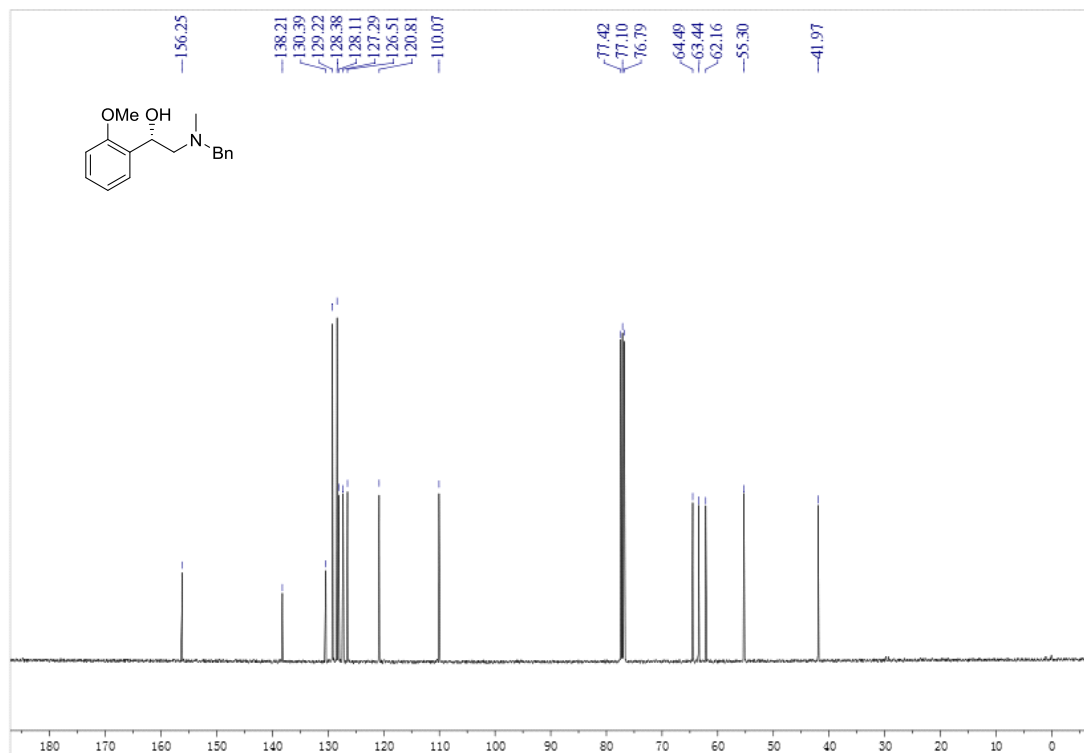
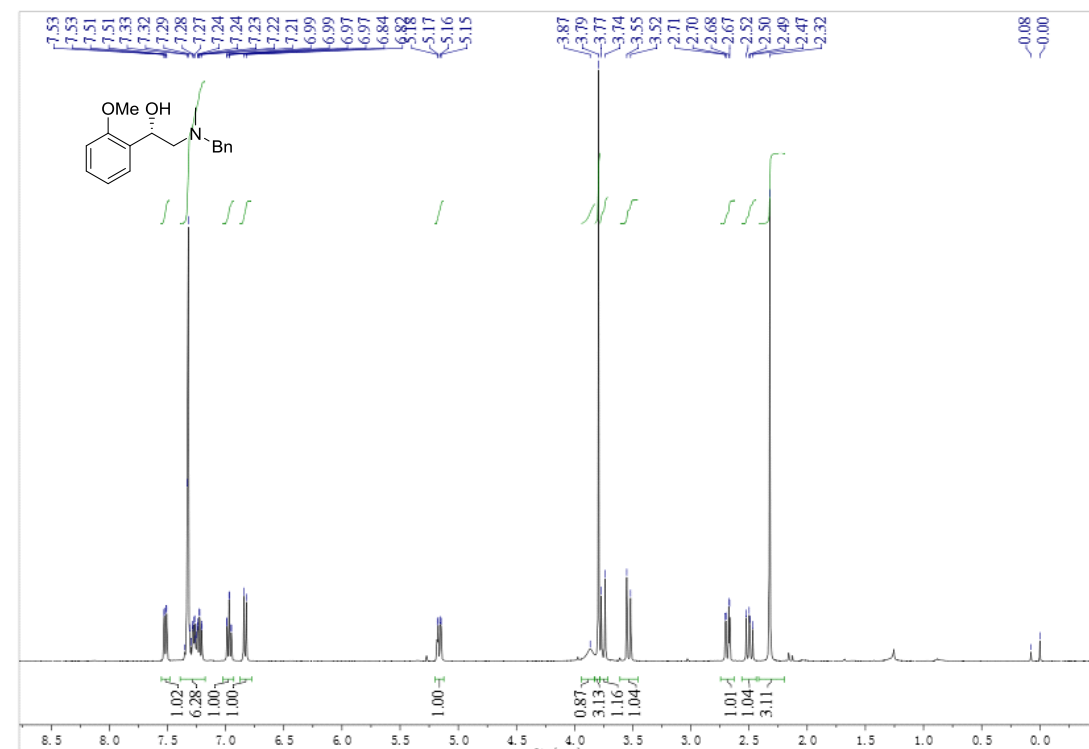
(S)-2-(Benzyl(methyl)amino)-1-(3-methylphenyl)ethanol (2b)



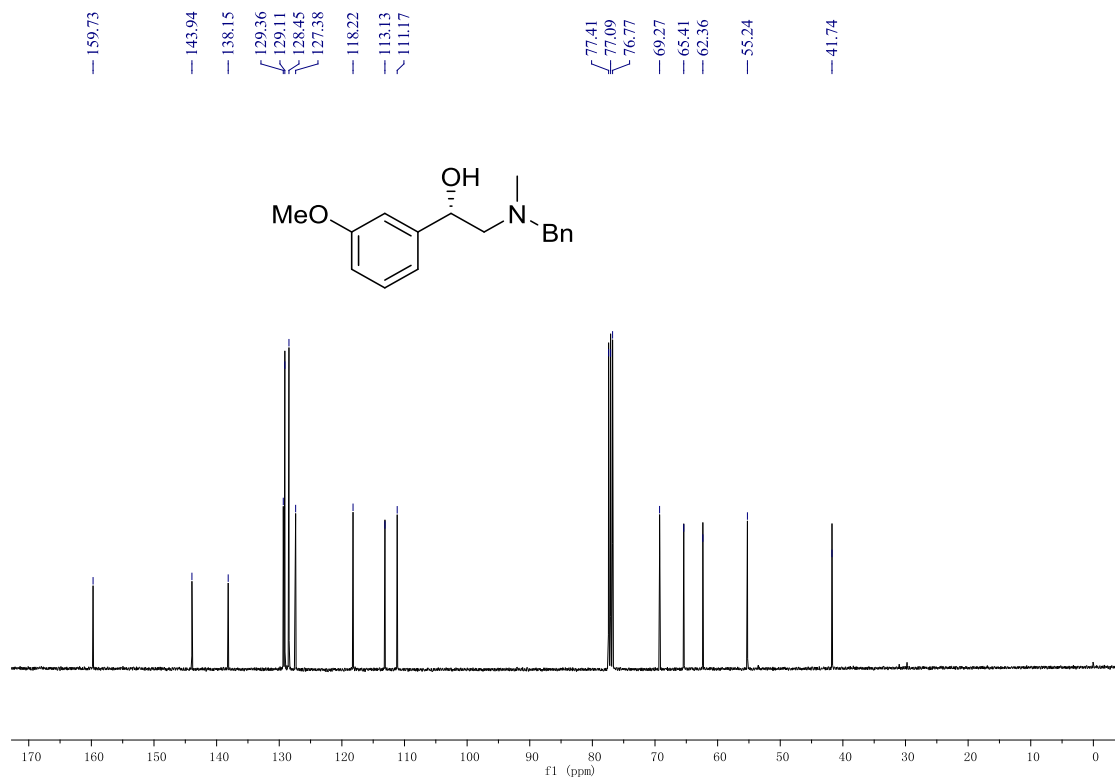
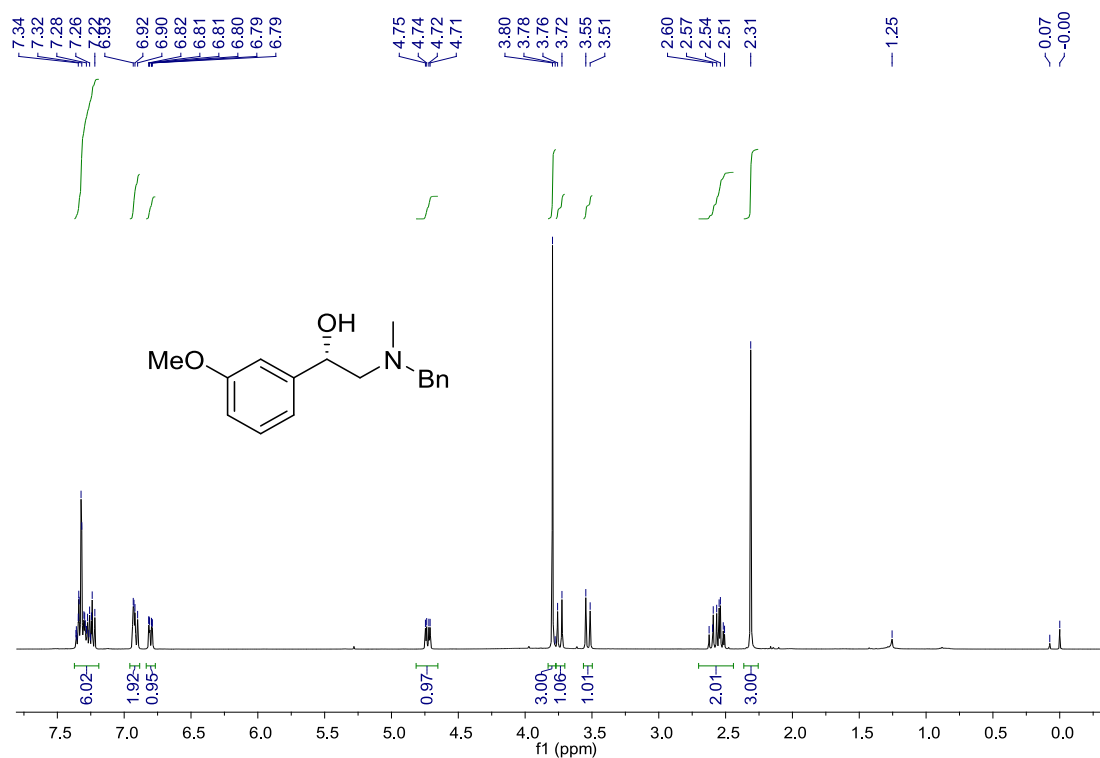
(S)-2-(Benzyl(methyl)amino)-1-(4-methylphenyl)ethanol(2c)



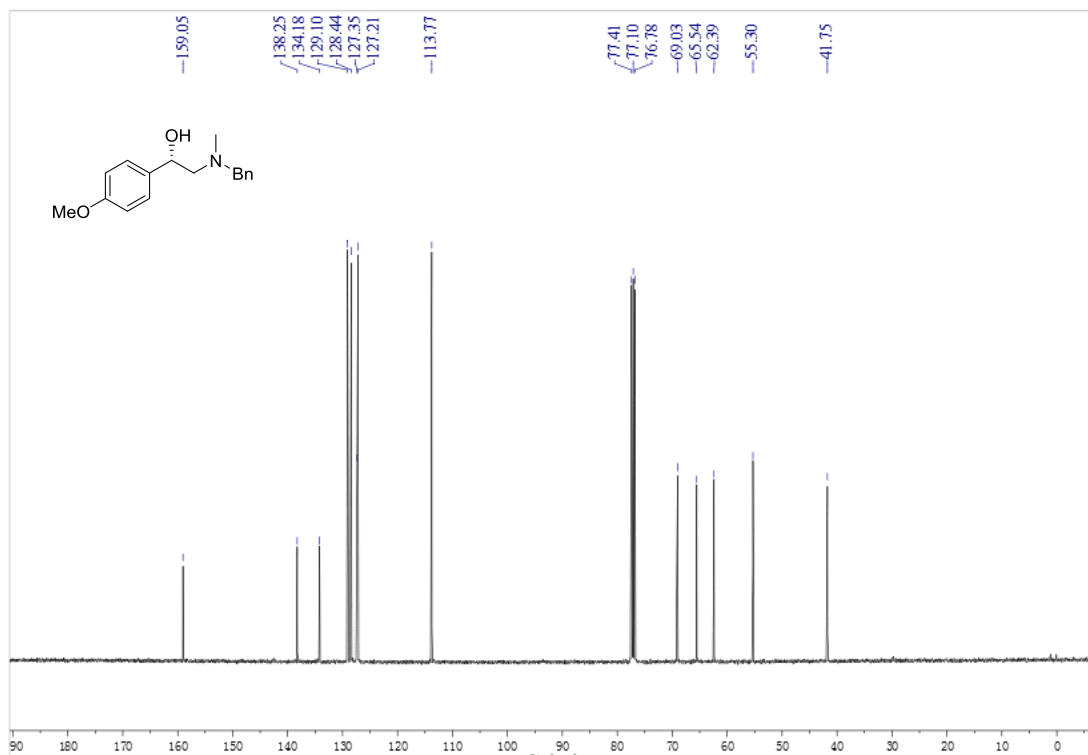
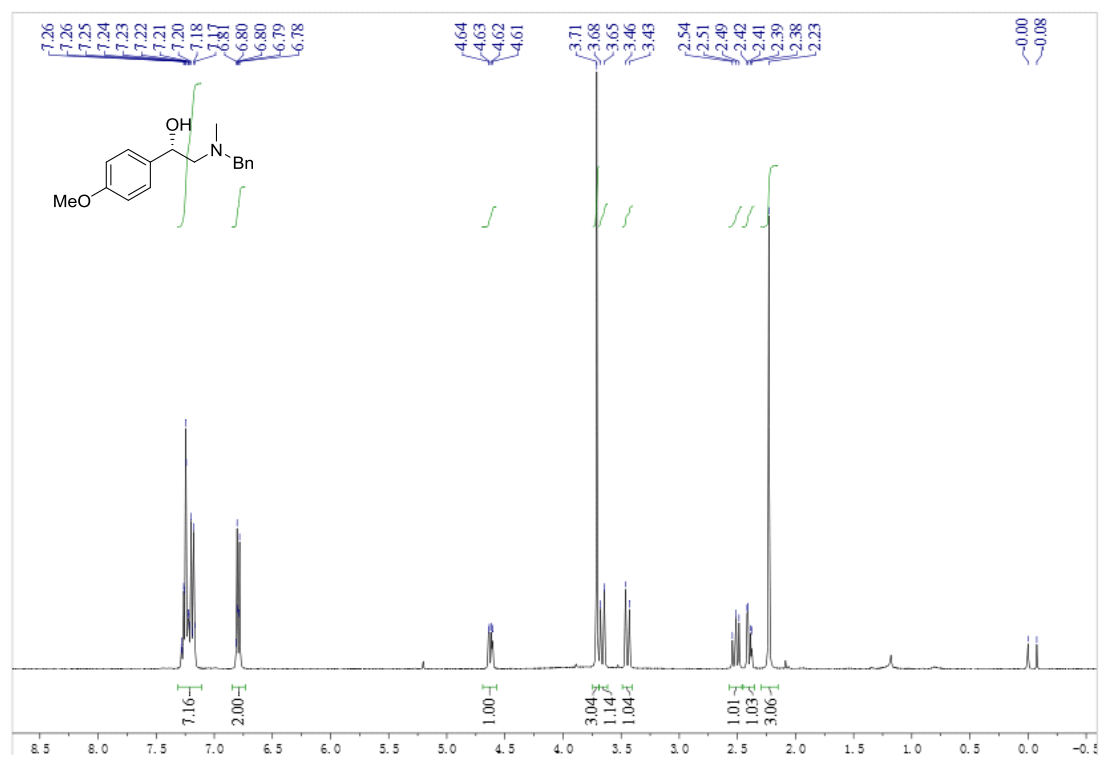
(S)-2-(Benzyl(methyl)amino)-1-(2-methoxyphenyl)ethanol (2d)



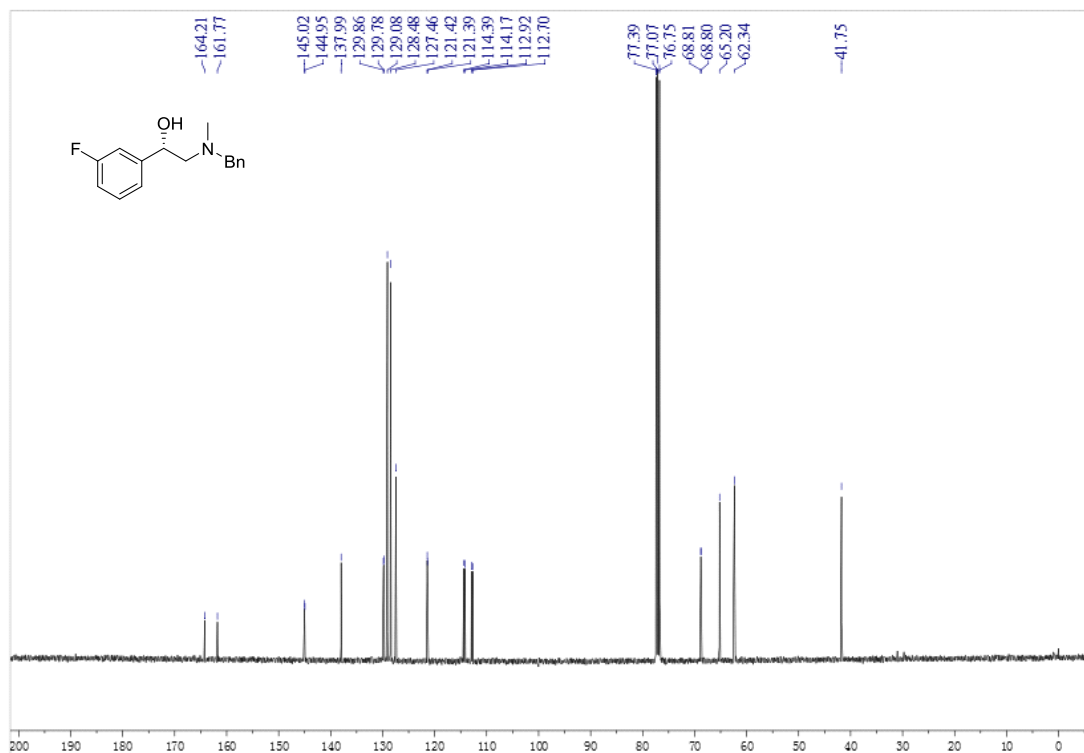
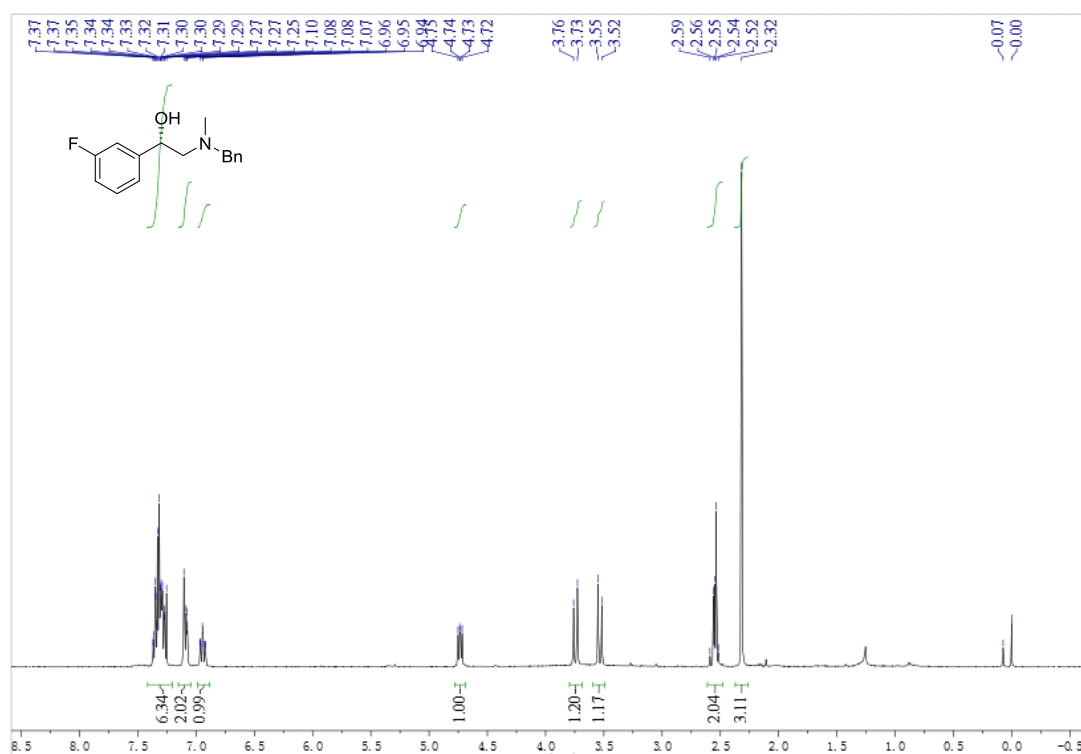
(S)-2-(Benzyl(methyl)amino)-1-(3-methoxyphenyl)ethanol (2e)



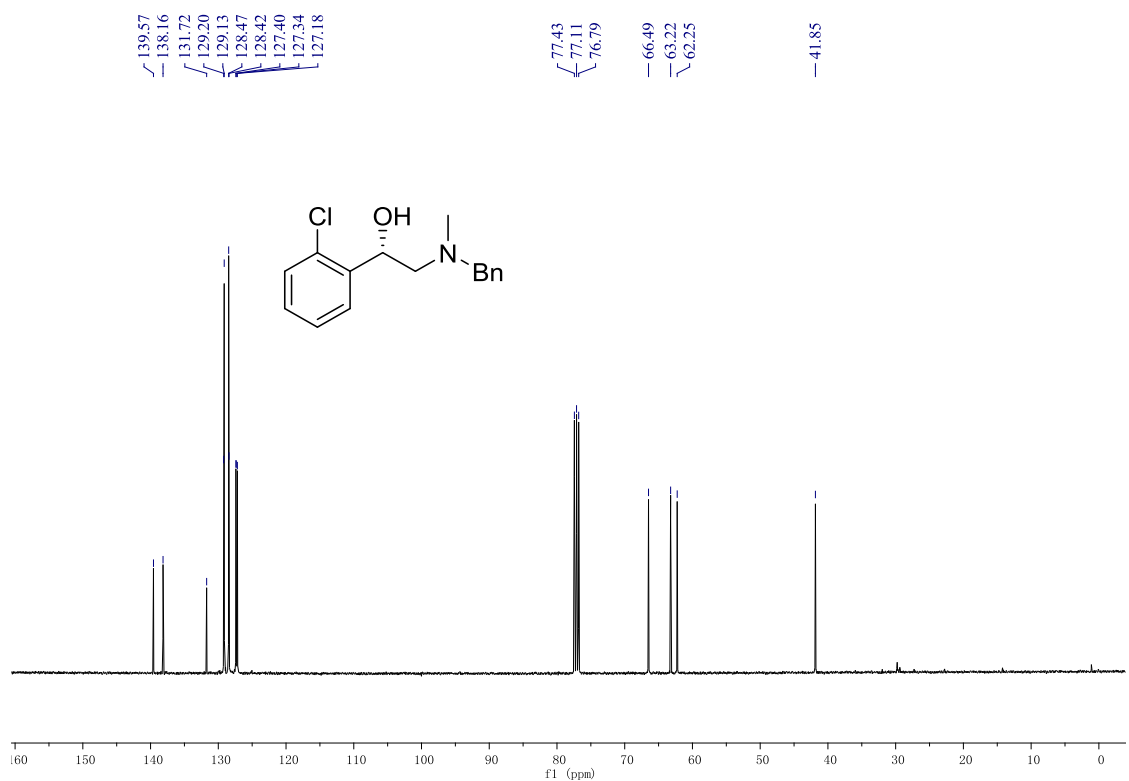
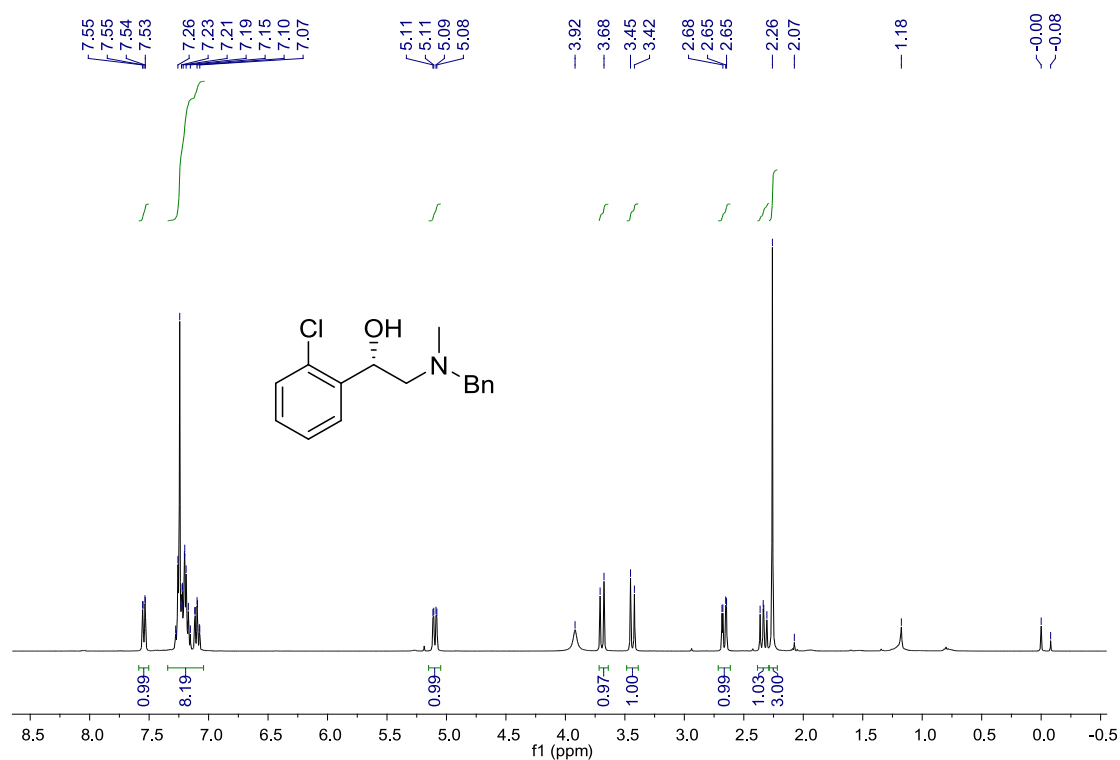
(S)-2-(Benzyl(methyl)amino)-1-(4-methoxyphenyl)ethanol (2f)



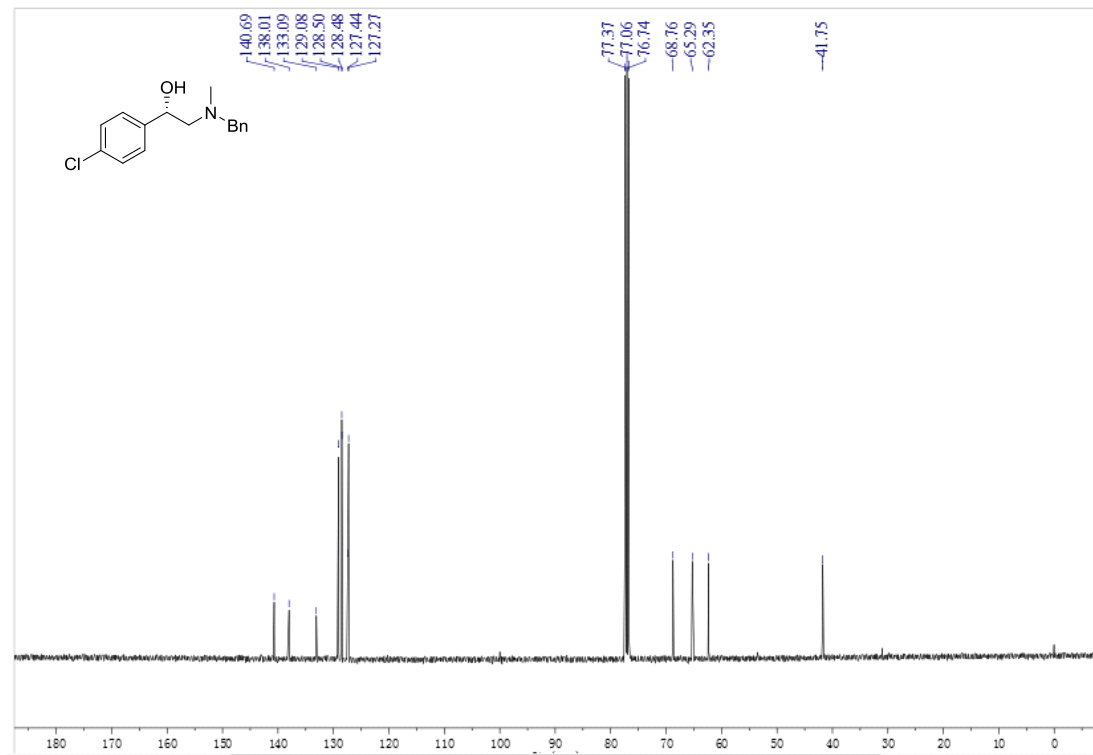
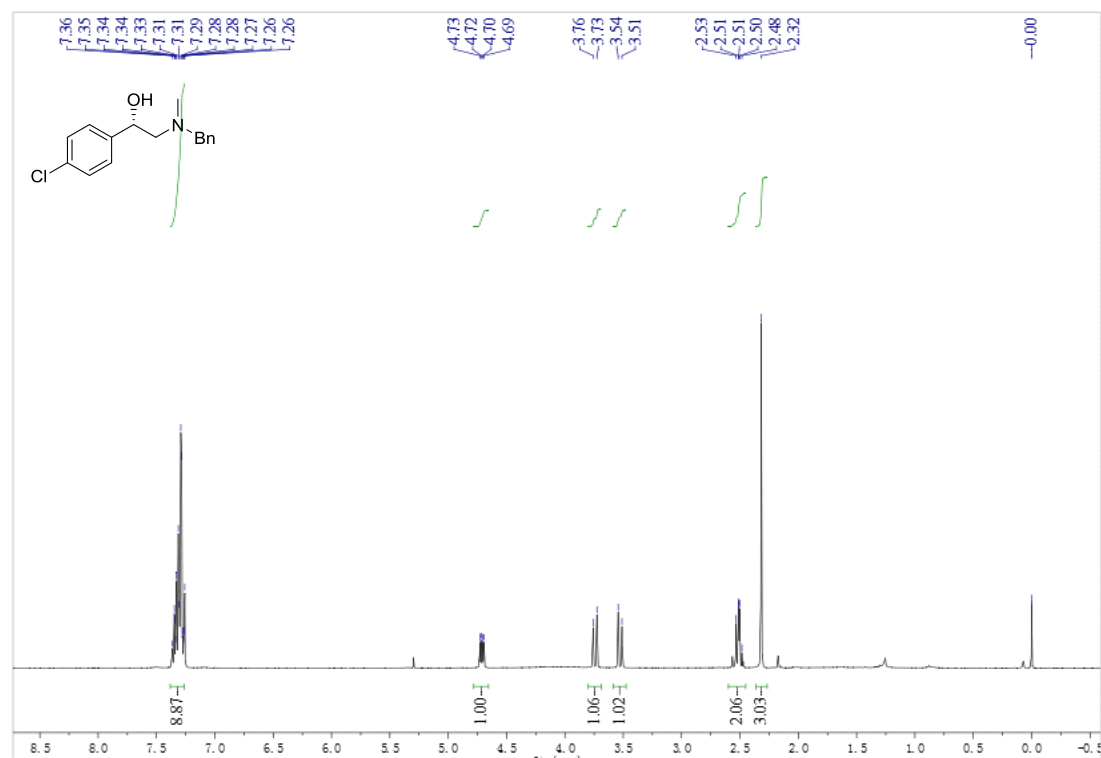
(S)-2-(benzyl(methyl)amino)-1-(3-fluorophenyl)ethanol (2g)



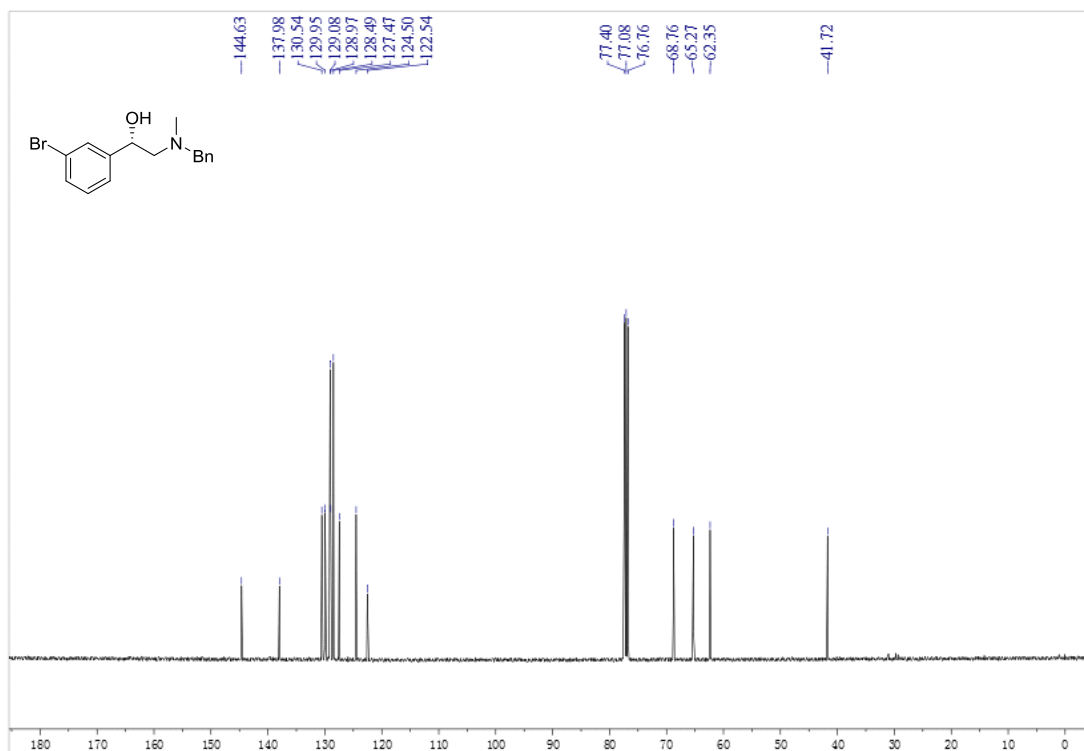
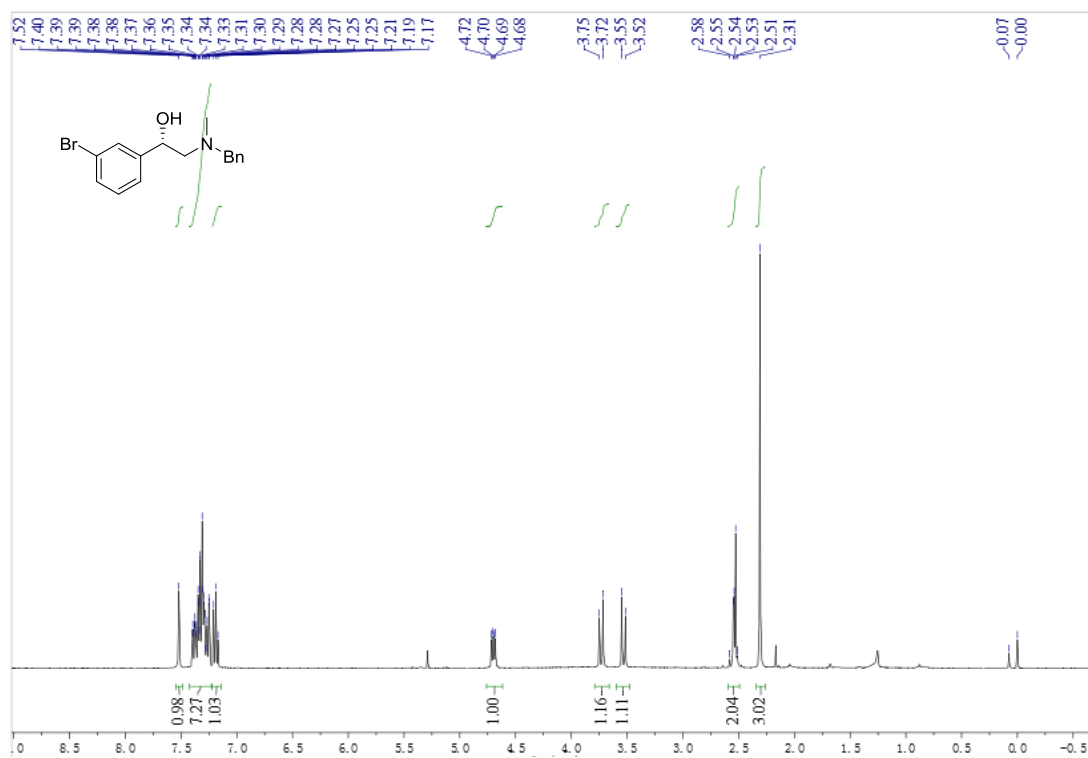
(S)-2-(Benzyl(methyl)amino)-1-(2-chlorophenyl)ethanol (2h)



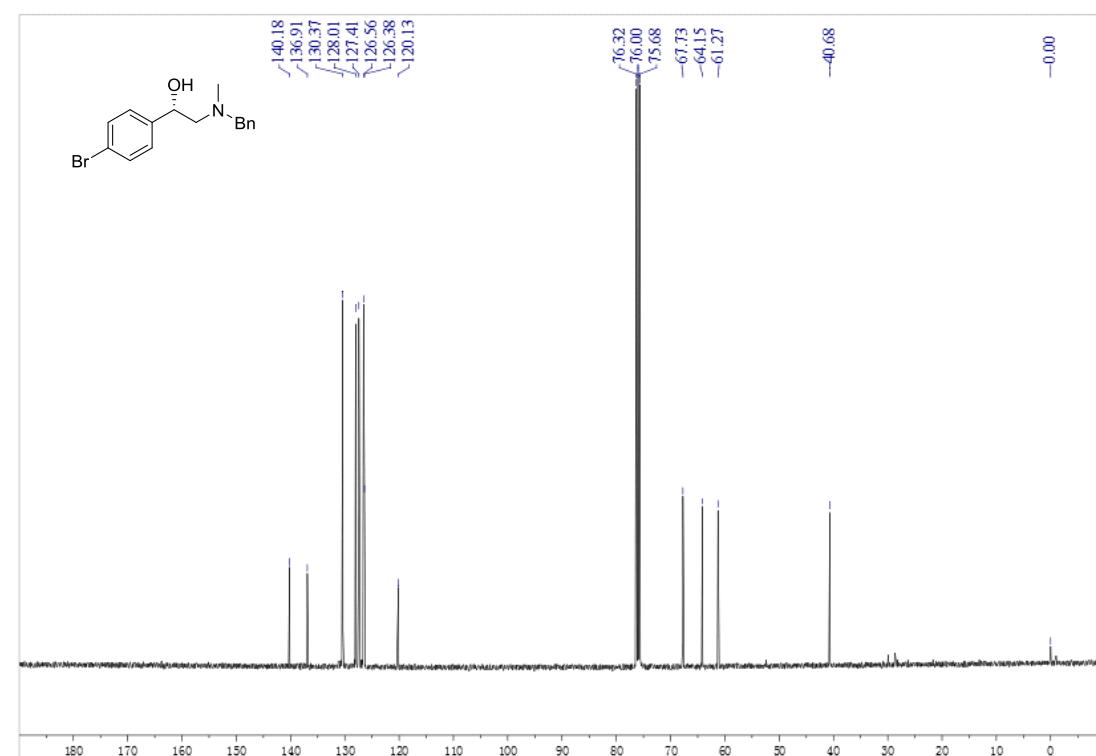
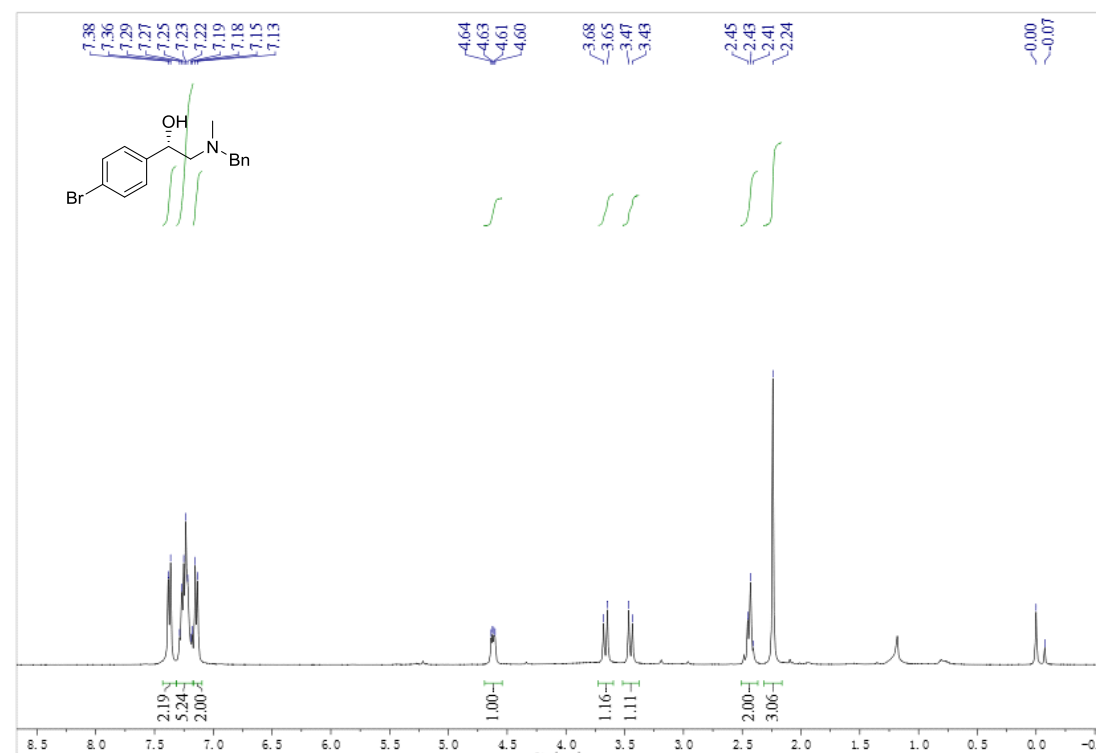
(S)-2-(Benzyl(methyl)amino)-1-(4-chlorophenyl)ethanol (2i)



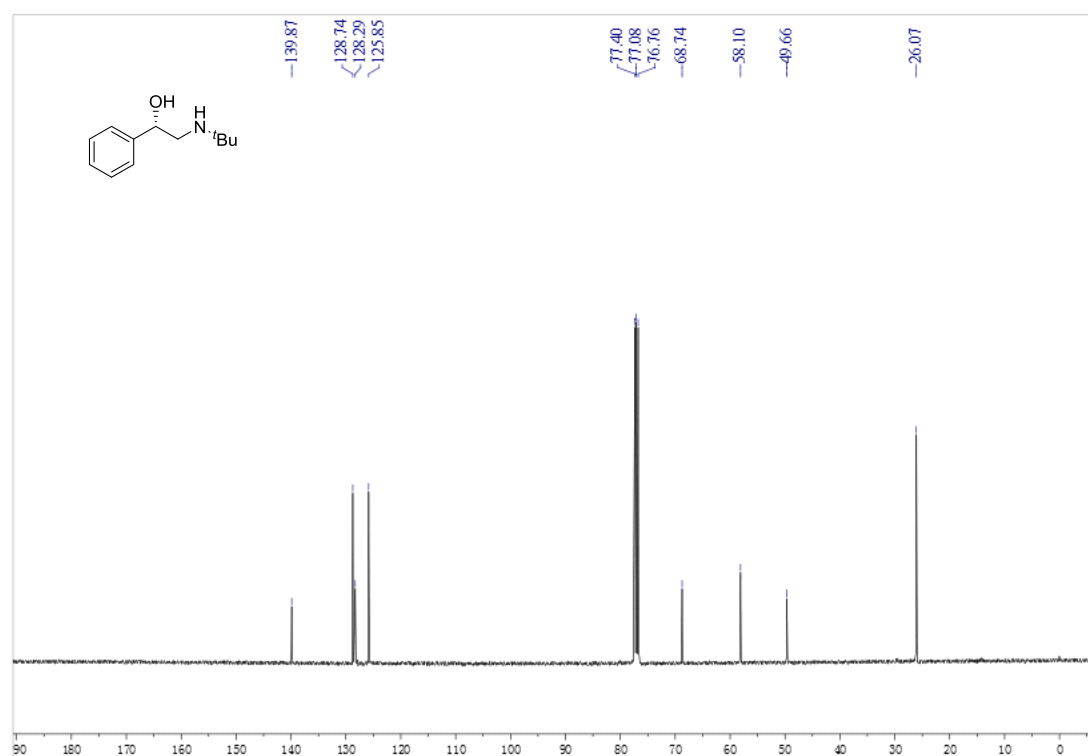
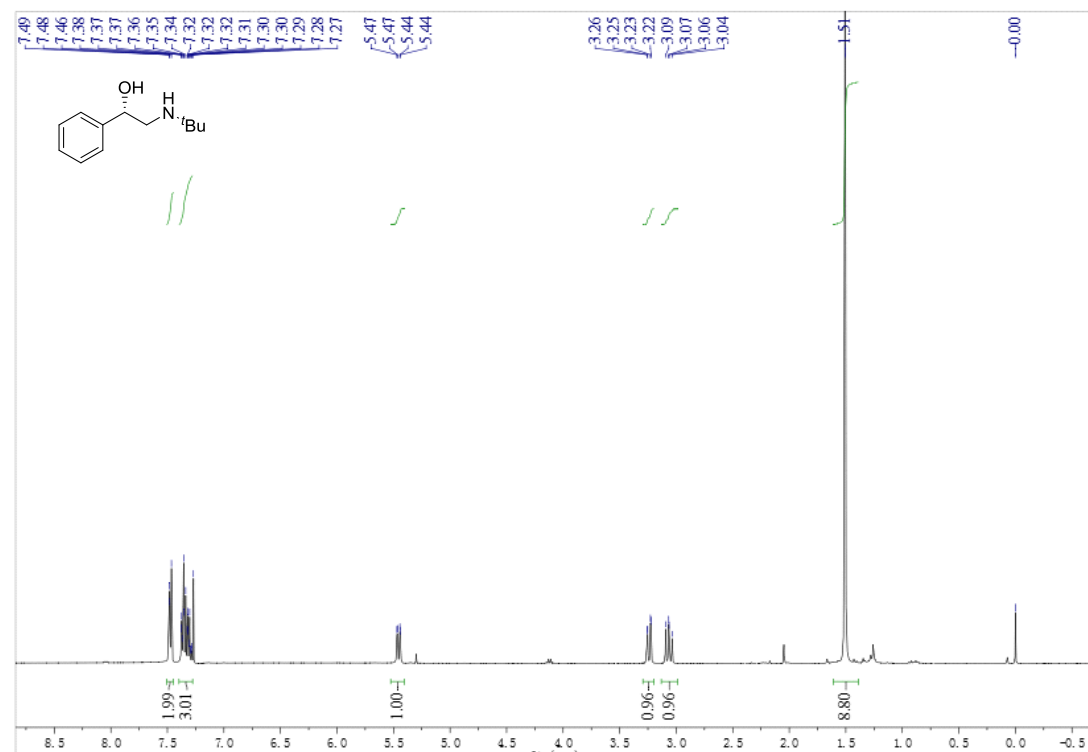
(S)-2-(Benzyl(methyl)amino)-1-(3-bromophenyl)ethanol (2j)



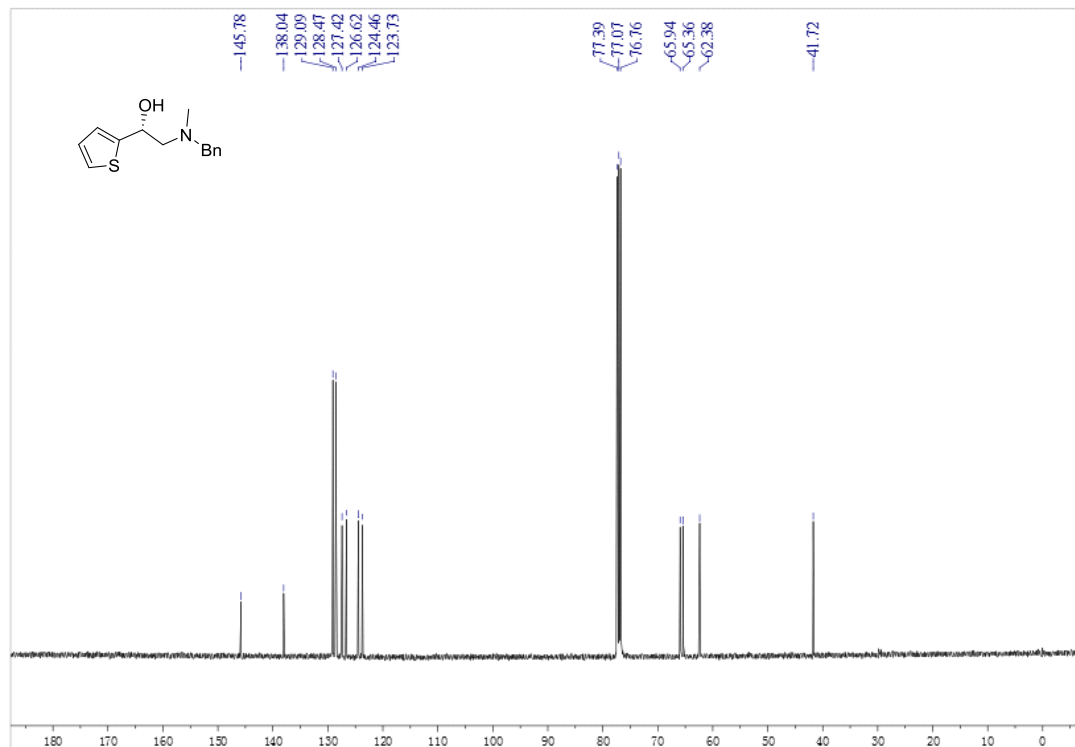
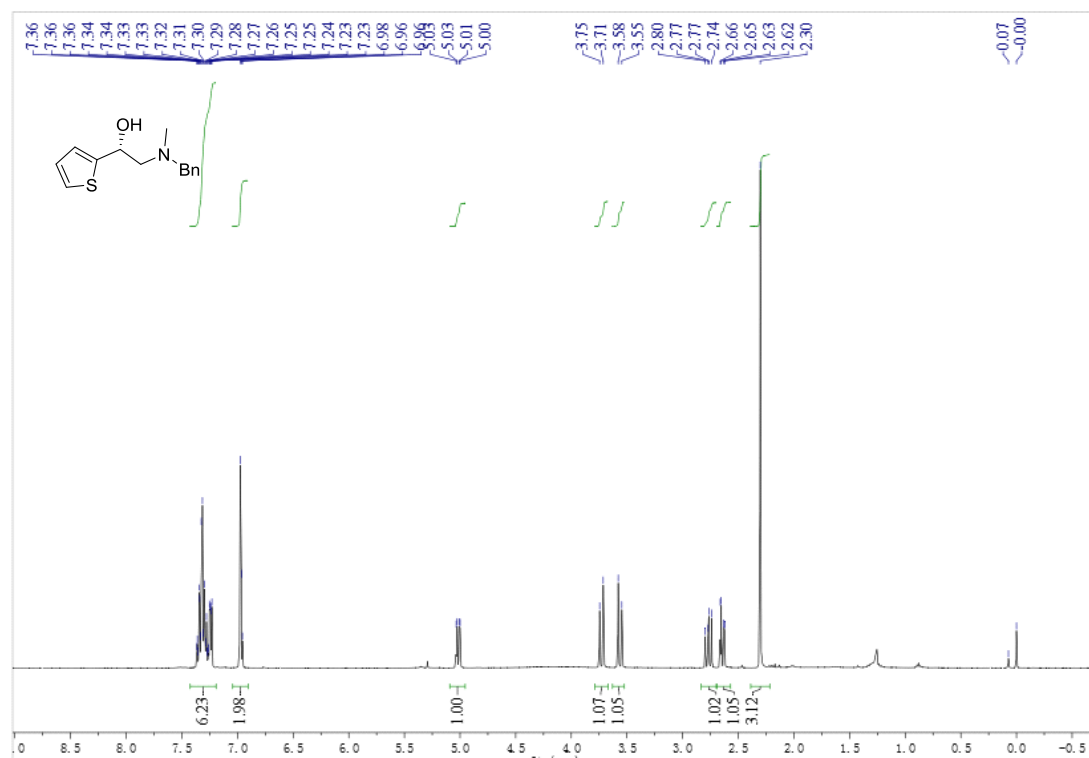
(S)-2-(Benzyl(methyl)amino)-1-(4-bromophenyl)ethanol (2k)



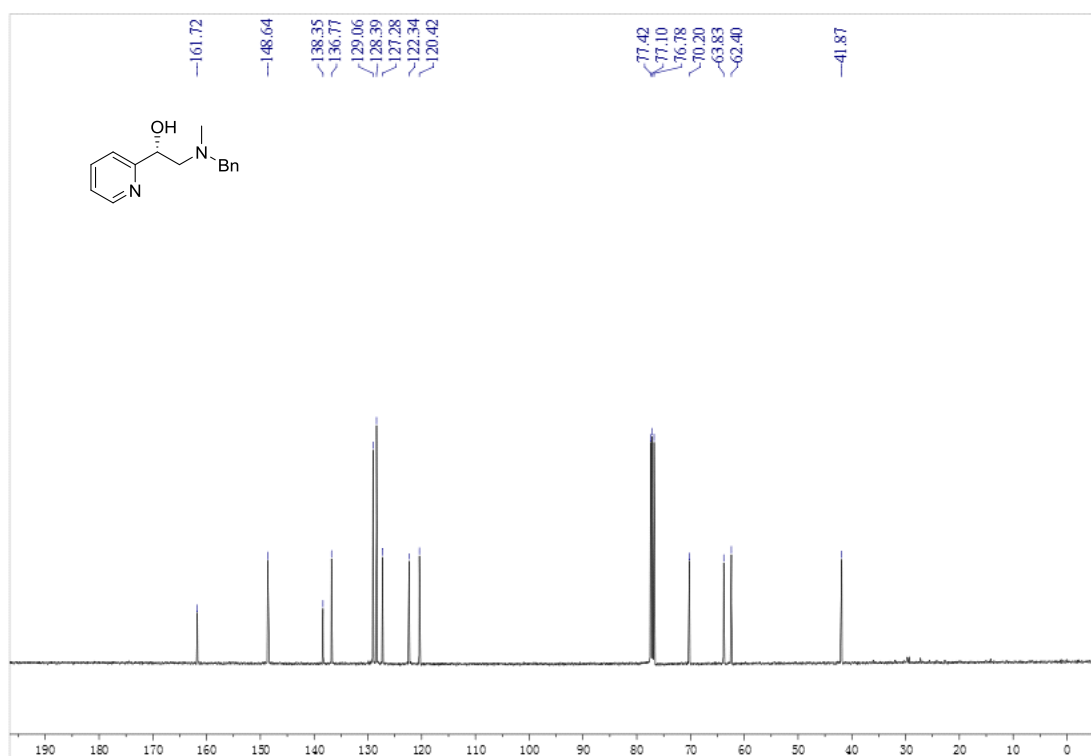
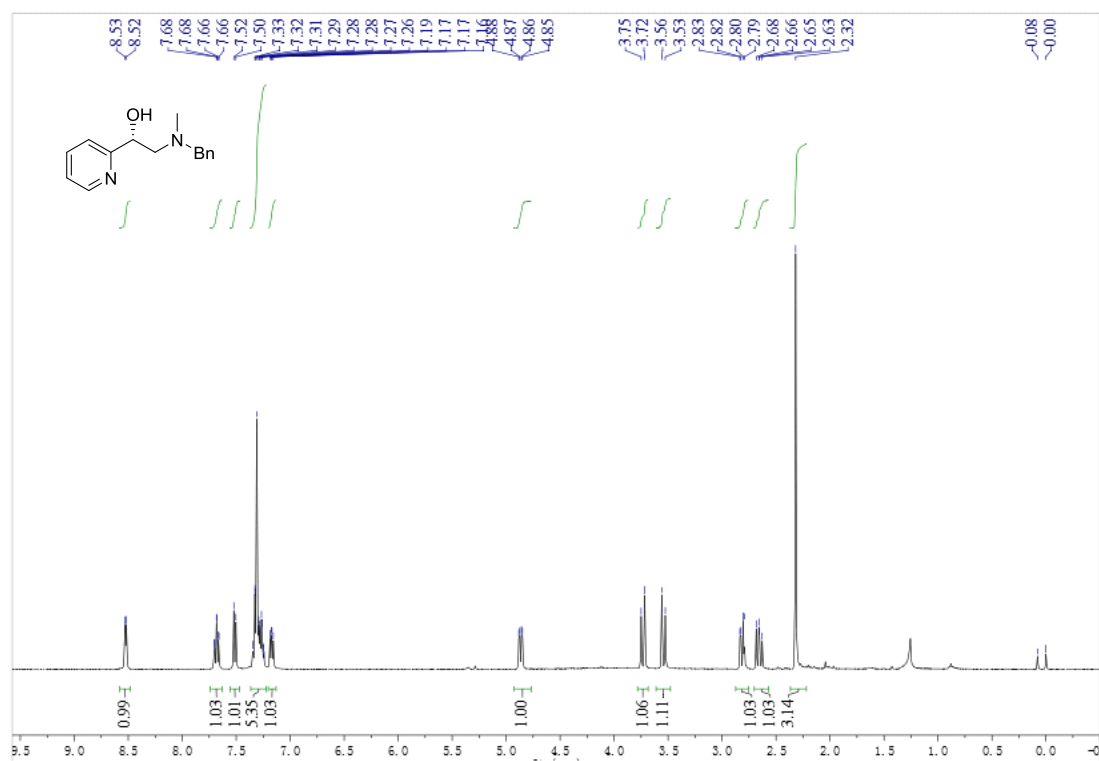
(S)-2-(tert-butylamino)-1-phenylethanol (2l)



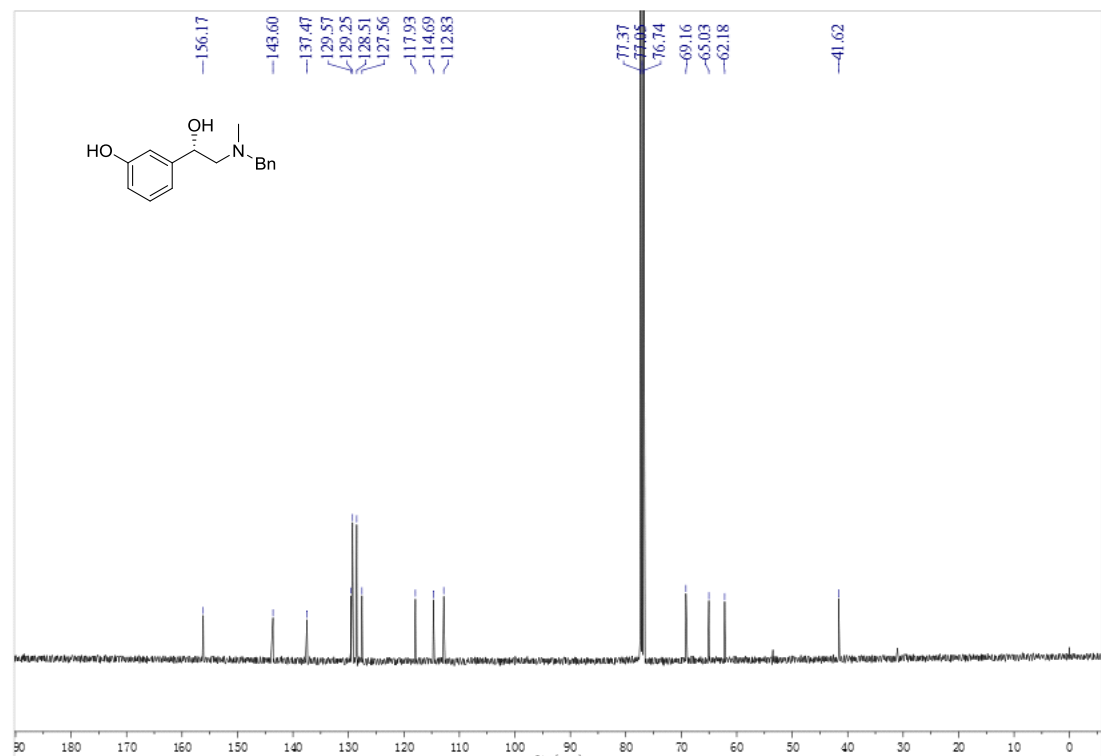
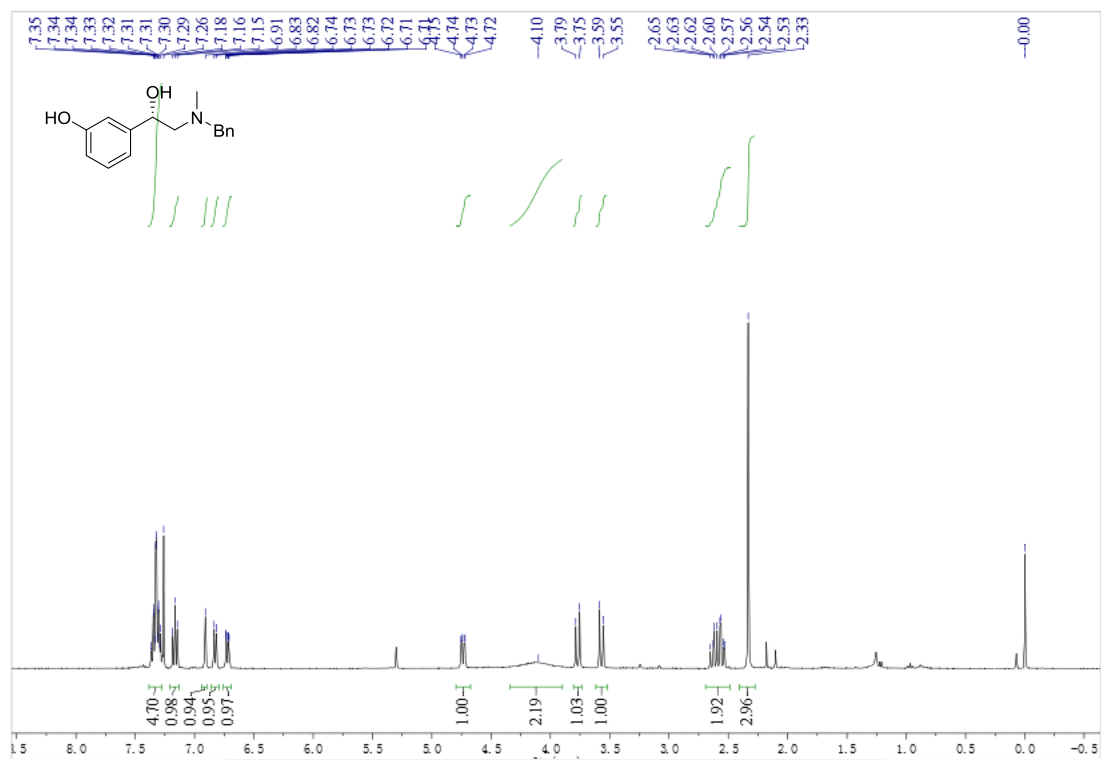
(R)-2-(benzyl(methyl)amino)-1-(thiophen-2-yl)ethanol (2m)



(R)-2-(benzyl(methyl)amino)-1-(pyridin-2-yl)ethanol (2n)



(S)-3-(2-(benzyl(methyl)amino)-1-hydroxyethyl)phenol (2o)

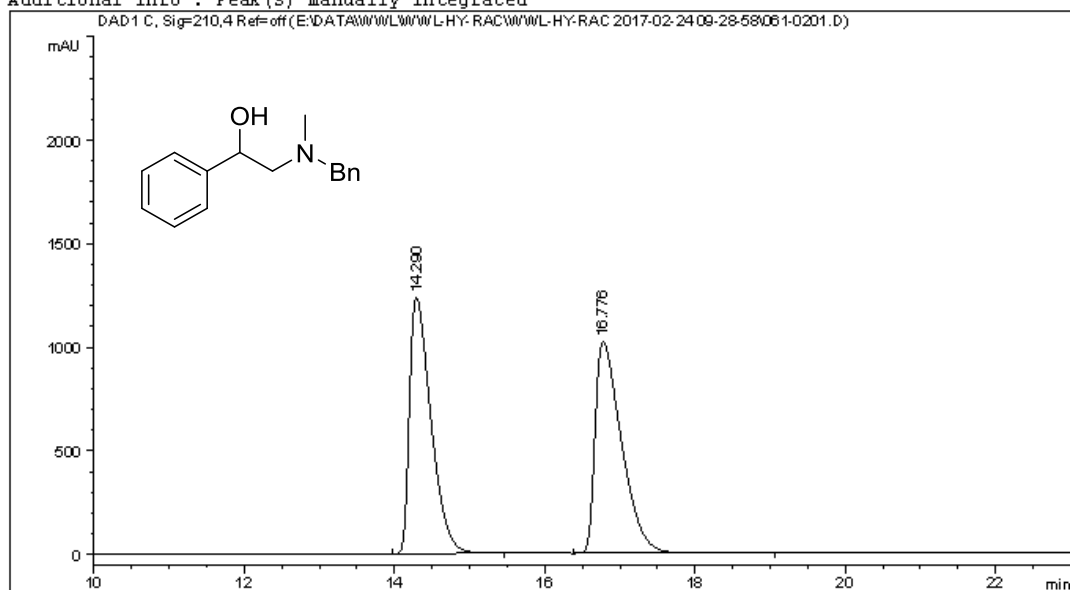


6. HPLC spectra

(S)-2-(Benzyl(methyl)amino)-1-phenylethanol (2a)

Data File E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\061-0201.D
Sample Name: PH-RAC

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Acq. Operator   : SYSTEM                      Seq. Line :    2
Acq. Instrument : 1260HPLC-DAD                Location  : Vial 61
Injection Date  : 2/24/2017 9:42:04 AM       Inj       :    1
                                           Inj Volume: 2.000 µl
Acq. Method     : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\DAD-0J(1-6)-95-5-
                  LML-2UL-ALL-30MIN.M
Last changed    : 2/24/2017 9:28:58 AM by SYSTEM
Analysis Method : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\DAD-0J(1-6)-95-5-
                  LML-2UL-ALL-30MIN.M (Sequence Method)
Last changed    : 2/24/2017 4:36:06 PM by SYSTEM
                  (modified after loading)
Additional Info : Peak(s) manually integrated
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Area Percent Report

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Sorted By      :      Signal
Multiplier     :      1.0000
Dilution       :      1.0000
Do not use Multiplier & Dilution Factor with ISTDs
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Signal 1: DAD1 C, Sig=210,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	14.290	BB	0.3052	2.45106e4	1236.98792	49.5107
2	16.776	BB	0.3688	2.49950e4	1024.02173	50.4893

Totals : 4.95056e4 2261.00964

*** End of Report ***

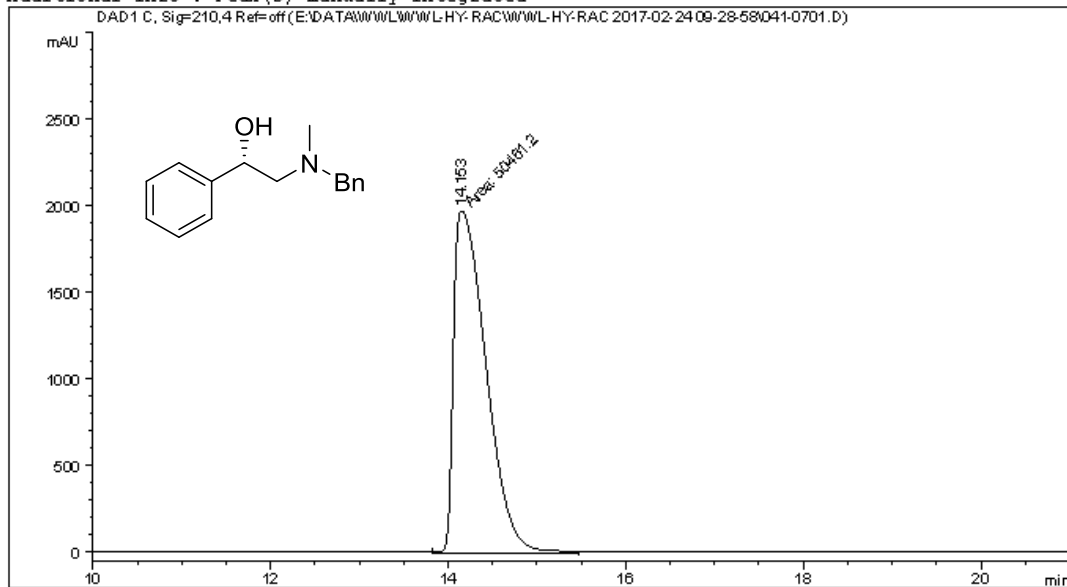
Data File E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\041-0701.D
 Sample Name: PH-EE

```

=====
Acq. Operator   : SYSTEM                      Seq. Line :    7
Acq. Instrument : 1260HPLC-DAD                Location  : Vial 41
Injection Date  : 2/24/2017 12:16:38 PM      Inj       :    1
                                           Inj Volume: 2.000 µl

Acq. Method     : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\DAD-0J(1-6)-95-5-
                  1ML-2UL-ALL-30MIN.M
Last changed    : 2/24/2017 9:28:58 AM by SYSTEM
Analysis Method : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\DAD-0J(1-6)-95-5-
                  1ML-2UL-ALL-30MIN.M (Sequence Method)
Last changed    : 2/24/2017 4:38:07 PM by SYSTEM
                  (modified after loading)
  
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Additional Info : Peak(s) manually integrated



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 Area Percent Report
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Sorted By      : Signal
Multiplier     : 1.0000
Dilution       : 1.0000
Do not use Multiplier & Dilution Factor with ISTDs
  
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Signal 1: DAD1 C, Sig=210,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	14.153	MM	0.4254	5.04612e4	1977.12036	100.0000

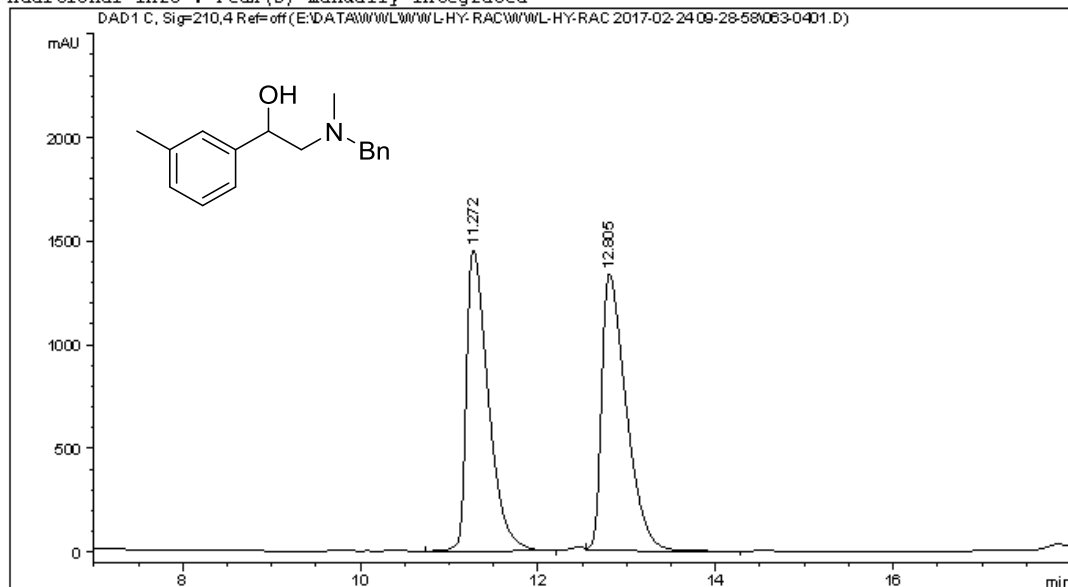
Totals : 5.04612e4 1977.12036

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 *** End of Report ***

(S)-2-(Benzyl(methyl)amino)-1-(3-methylphenyl)ethanol (2b)

Data File E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\063-0401.D
Sample Name: M-ME-RAC

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Acq. Operator   : SYSTEM                      Seq. Line :    4
Acq. Instrument : 1260HPLC-DAD                Location  : Vial 63
Injection Date  : 2/24/2017 10:43:54 AM      Inj       :    1
                                           Inj Volume: 2.000 µl
Acq. Method     : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\DAD-0J(1-6)-95-5-
                  LML-2UL-ALL-30MIN.M
Last changed    : 2/24/2017 9:28:58 AM by SYSTEM
Analysis Method : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\DAD-0J(1-6)-95-5-
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Last changed    : 2/24/2017 4:22:58 PM by SYSTEM
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Additional Info  : Peak(s) manually integrated
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Area Percent Report

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Sorted By      :      Signal
Multiplier     :      1.0000
Dilution       :      1.0000
Do not use Multiplier & Dilution Factor with ISTDs
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Signal 1: DAD1 C, Sig=210,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	11.272	BB	0.2618	2.48888e4	1453.32751	49.2322
2	12.805	VB	0.2900	2.56651e4	1336.59631	50.7678

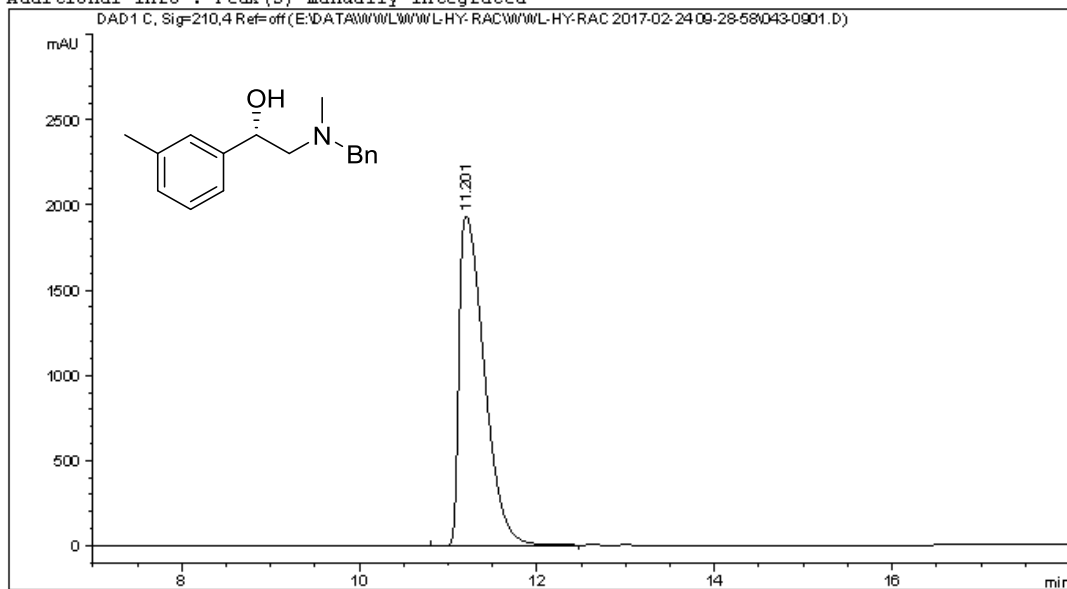
Totals : 5.05540e4 2789.92383

*** End of Report ***

Data File E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\043-0901.D
Sample Name: M-ME-EE

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Acq. Operator   : SYSTEM                      Seq. Line :    9
Acq. Instrument : 1260HPLC-DAD                Location  : Vial 43
Injection Date  : 2/24/2017 1:18:24 PM        Inj       :    1
                                           Inj Volume: 2.000 µl

Acq. Method     : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\DAD-0J(1-6)-95-5-
                  LML-2UL-ALL-30MIN.M
Last changed    : 2/24/2017 9:28:58 AM by SYSTEM
Analysis Method : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\DAD-0J(1-6)-95-5-
                  LML-2UL-ALL-30MIN.M (Sequence Method)
Last changed    : 2/24/2017 4:24:25 PM by SYSTEM
                  (modified after loading)
Additional Info : Peak(s) manually integrated
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Area Percent Report
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Sorted By : Signal
Multiplier : 1.0000
Dilution : 1.0000
Do not use Multiplier & Dilution Factor with ISTDs

Signal 1: DAD1 C, Sig=210,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	11.201	BB	0.3043	3.78525e4	1934.52808	100.0000

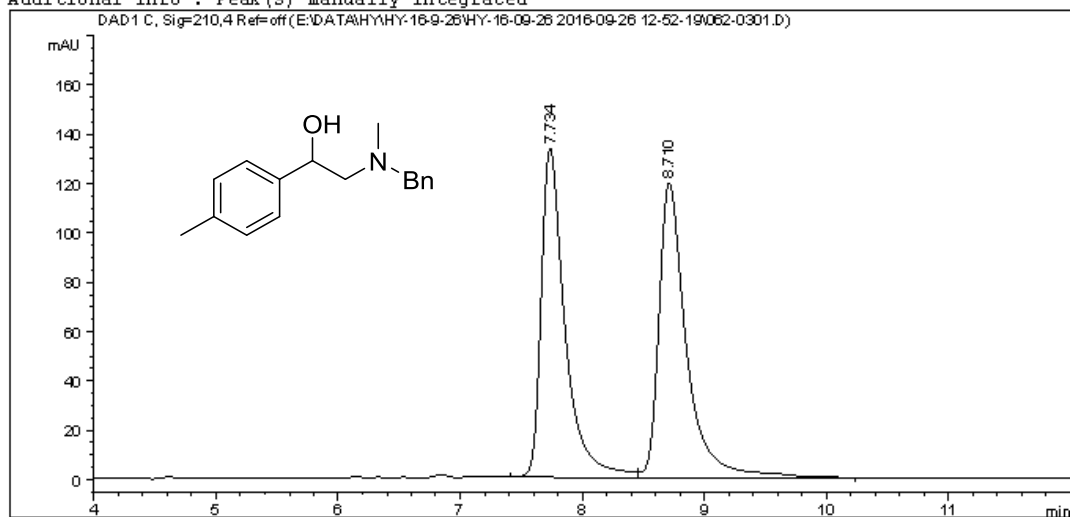
Totals : 3.78525e4 1934.52808

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*** End of Report ***

(S)-2-(Benzyl(methyl)amino)-1-(4-methylphenyl)ethanol (2c)

Data File E:\DATA\HY\HY-16-9-26\HY-16-09-26 2016-09-26 12-52-19\062-0301.D
Sample Name: HY-598

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Acq. Operator   : SYSTEM                      Seq. Line :    3
Acq. Instrument : 1260HPLC-DAD                Location  : Vial 62
Injection Date  : 9/26/2016 1:20:05 PM        Inj       :    1
                                           Inj Volume: 1.000 µl
Acq. Method     : E:\DATA\HY\HY-16-9-26\HY-16-09-26 2016-09-26 12-52-19\DAD-0J(1-6)-85-15-
                  1.OML-ALLNM-15MIN.M
Last changed    : 9/26/2016 12:52:20 PM by SYSTEM
Analysis Method : E:\DATA\HY\HY-16-9-26\HY-16-09-26 2016-09-26 12-52-19\DAD-0J(1-6)-85-15-
                  1.OML-ALLNM-15MIN.M (Sequence Method)
Last changed    : 2/9/2017 8:40:58 PM by SYSTEM
                  (modified after loading)
Additional Info  : Peak(s) manually integrated
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Area Percent Report

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Sorted By      : Signal
Multiplier     : 1.0000
Dilution       : 1.0000
Do not use Multiplier & Dilution Factor with ISTDs
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Signal 1: DAD1 C, Sig=210,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
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2	8.710	VB	0.2310	1875.80273	119.59895	50.8968

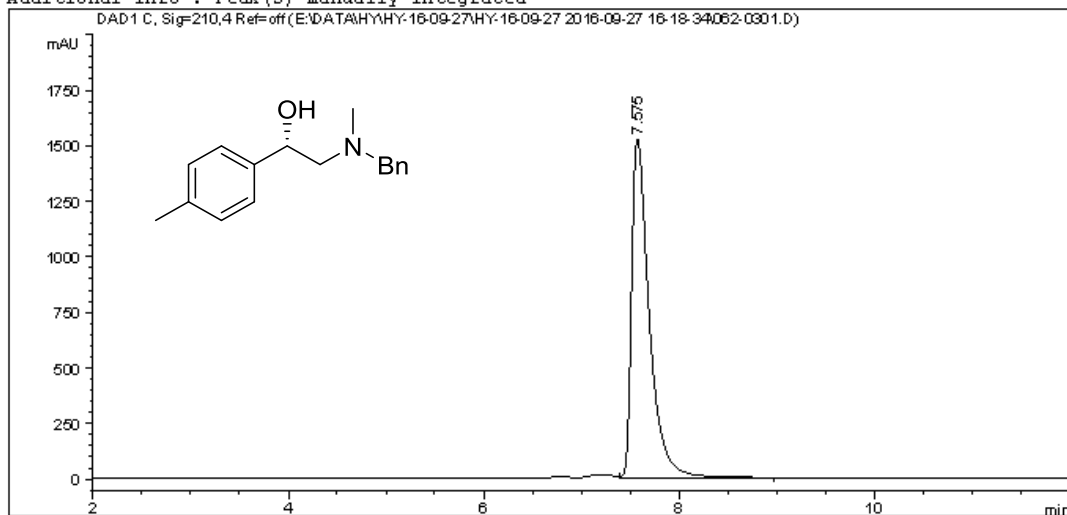
Totals : 3685.50354 253.08938

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=====
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Injection Date  : 9/27/2016 4:51:23 PM                 Inj       :    1
                                                    Inj Volume: 1.000 µl

Acq. Method     : E:\DATA\HY\HY-16-09-27\HY-16-09-27 2016-09-27 16-18-34\DAD-0J(1-6)-85-15
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Last changed    : 9/27/2016 4:18:35 PM by SYSTEM
Analysis Method : E:\DATA\HY\HY-16-09-27\HY-16-09-27 2016-09-27 16-18-34\DAD-0J(1-6)-85-15
                  -1.0ML-ALLNM-15MIN.M (Sequence Method)
Last changed    : 2/9/2017 8:43:44 PM by SYSTEM
                  (modified after loading)
Additional Info : Peak(s) manually integrated
  
```



=====
 Area Percent Report
 =====

```

Sorted By      :      Signal
Multiplier     :      1.0000
Dilution      :      1.0000
Do not use Multiplier & Dilution Factor with ISTDs
  
```

Signal 1: DAD1 C, Sig=210,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	7.575	VB	0.1889	1.92436e4	1528.72388	100.0000

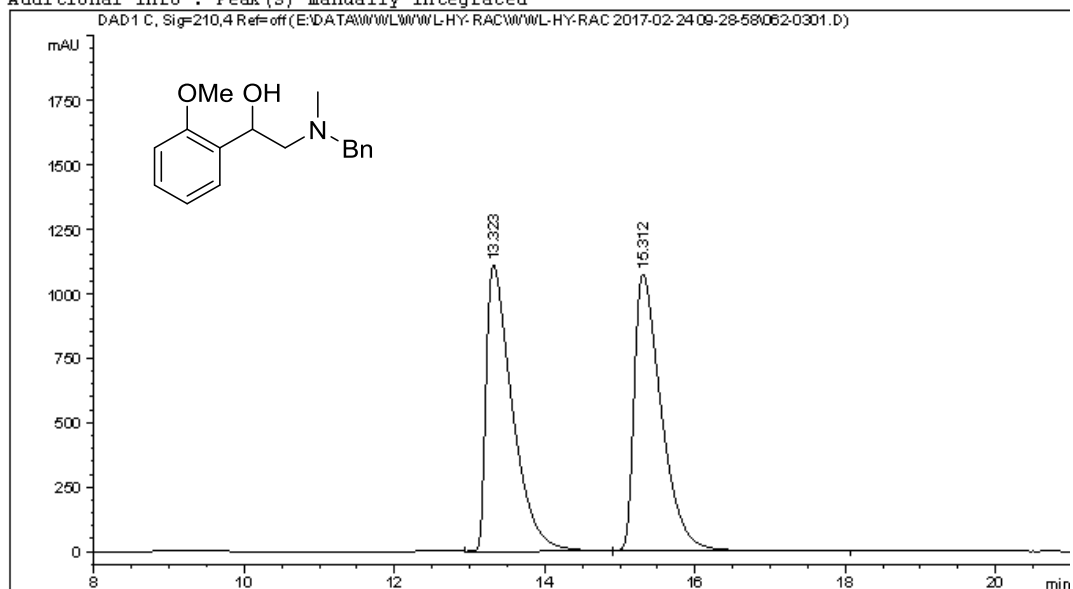
Totals : 1.92436e4 1528.72388

=====
 *** End of Report ***

(S)-2-(Benzyl(methyl)amino)-1-(2-methoxyphenyl)ethanol (2d)

Data File E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\062-0301.D
Sample Name: 0-ME0-RAC

```
=====
Acq. Operator   : SYSTEM                      Seq. Line :    3
Acq. Instrument : 1260HPLC-DAD                Location  : Vial 62
Injection Date  : 2/24/2017 10:12:59 AM      Inj       :    1
                                           Inj Volume: 2.000 µl
Acq. Method     : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\DAD-0J(1-6)-95-5-
                  LML-2UL-ALL-30MIN.M
Last changed    : 2/24/2017 9:28:58 AM by SYSTEM
Analysis Method : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\DAD-0J(1-6)-95-5-
                  LML-2UL-ALL-30MIN.M (Sequence Method)
Last changed    : 2/24/2017 4:26:10 PM by SYSTEM
                  (modified after loading)
Additional Info  : Peak(s) manually integrated
=====
```



Area Percent Report

```
Sorted By      :      Signal
Multiplier     :      1.0000
Dilution       :      1.0000
Do not use Multiplier & Dilution Factor with ISTDs
```

Signal 1: DAD1 C, Sig=210,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	13.323	BV	0.3605	2.68774e4	1109.82776	49.8642
2	15.312	VB	0.3791	2.70238e4	1075.81323	50.1358

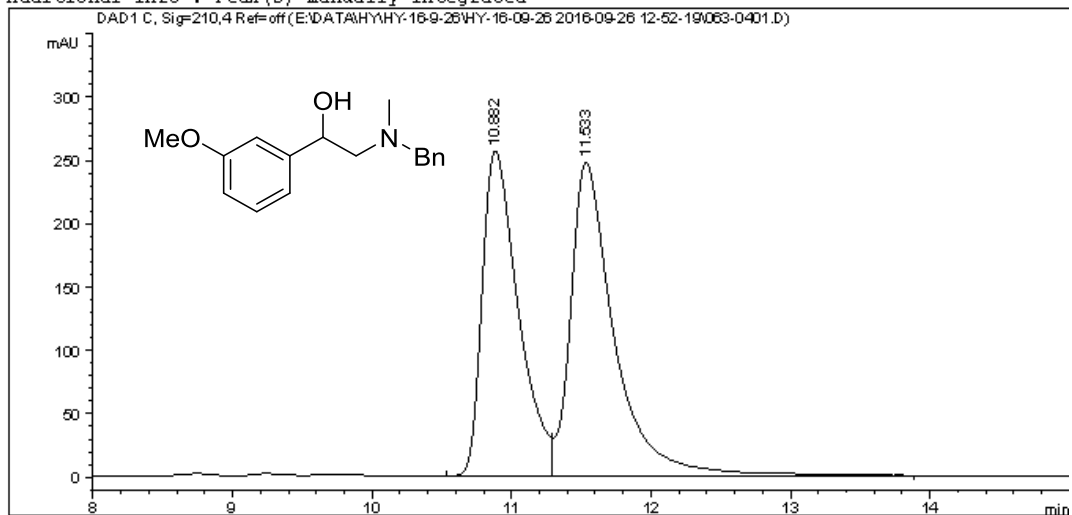
Totals : 5.39012e4 2185.64099

*** End of Report ***

(S)-2-(Benzyl(methyl)amino)-1-(3-methoxyphenyl)ethanol (2e)

Data File E:\DATA\HY\HY-16-9-26\HY-16-09-26 2016-09-26 12-52-19\063-0401.D
Sample Name: HY-599

```
=====
Acq. Operator   : SYSTEM                      Seq. Line :    4
Acq. Instrument : 1260HPLC-DAD                Location  : Vial 63
Injection Date  : 9/26/2016 1:35:58 PM        Inj       :    1
                                           Inj Volume: 1.000 µl
Acq. Method     : E:\DATA\HY\HY-16-9-26\HY-16-09-26 2016-09-26 12-52-19\DAD-0J(1-6)-85-15-
                  1.OML-ALLNM-15MIN.M
Last changed    : 9/26/2016 12:52:20 PM by SYSTEM
Analysis Method : E:\DATA\HY\HY-16-9-26\HY-16-09-26 2016-09-26 12-52-19\DAD-0J(1-6)-85-15-
                  1.OML-ALLNM-15MIN.M (Sequence Method)
Last changed    : 2/9/2017 8:47:00 PM by SYSTEM
                  (modified after loading)
Additional Info  : Peak(s) manually integrated
=====
```



Area Percent Report

```
Sorted By      :      Signal
Multiplier     :      1.0000
Dilution      :      1.0000
Do not use Multiplier & Dilution Factor with ISTDs
```

Signal 1: DAD1 C, Sig=210,4 Ref=off

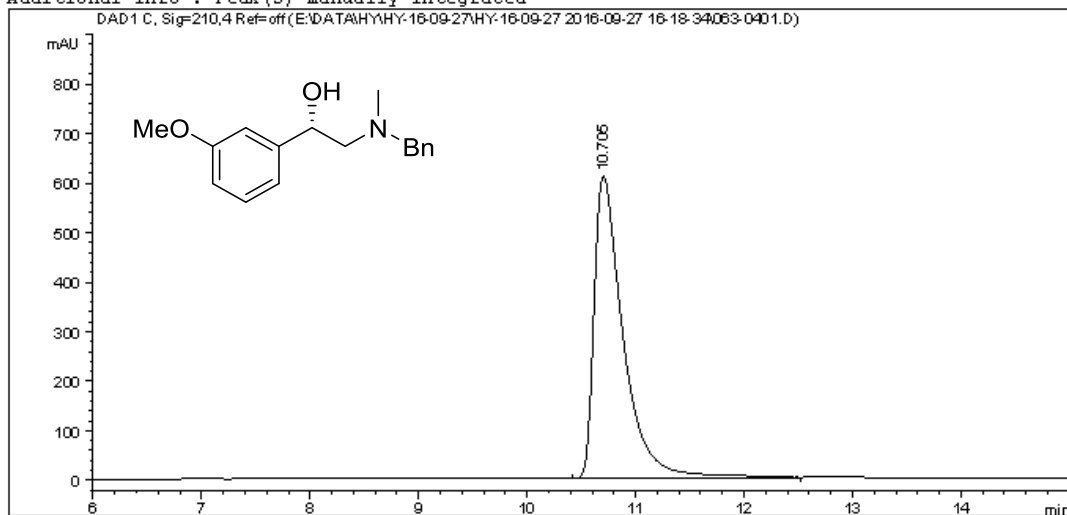
Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	10.882	BV	0.2700	4619.91064	256.70032	46.2962
2	11.533	VB	0.3163	5359.11670	247.79356	53.7038

Totals : 9979.02734 504.49388

*** End of Report ***

```

=====
Acq. Operator   : SYSTEM                               Seq. Line :    4
Acq. Instrument : 1260HPLC-DAD                         Location  : Vial 63
Injection Date  : 9/27/2016 5:07:19 PM                Inj       :    1
                                                    Inj Volume: 1.000 µl
Acq. Method     : E:\DATA\HY\HY-16-09-27\HY-16-09-27 2016-09-27 16-18-34\DAD-0J(1-6)-85-15
                  -1.0ML-ALLNM-15MIN.M
Last changed    : 9/27/2016 4:18:35 PM by SYSTEM
Analysis Method : E:\DATA\HY\HY-16-09-27\HY-16-09-27 2016-09-27 16-18-34\DAD-0J(1-6)-85-15
                  -1.0ML-ALLNM-15MIN.M (Sequence Method)
Last changed    : 2/9/2017 8:50:50 PM by SYSTEM
                  (modified after loading)
Additional Info  : Peak(s) manually integrated
  
```



=====
 Area Percent Report
 =====

```

Sorted By      :      Signal
Multiplier     :      1.0000
Dilution      :      1.0000
Do not use Multiplier & Dilution Factor with ISTDs
  
```

Signal 1: DAD1 C, Sig=210,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	10.705	BB	0.2754	1.13701e4	610.29456	100.0000

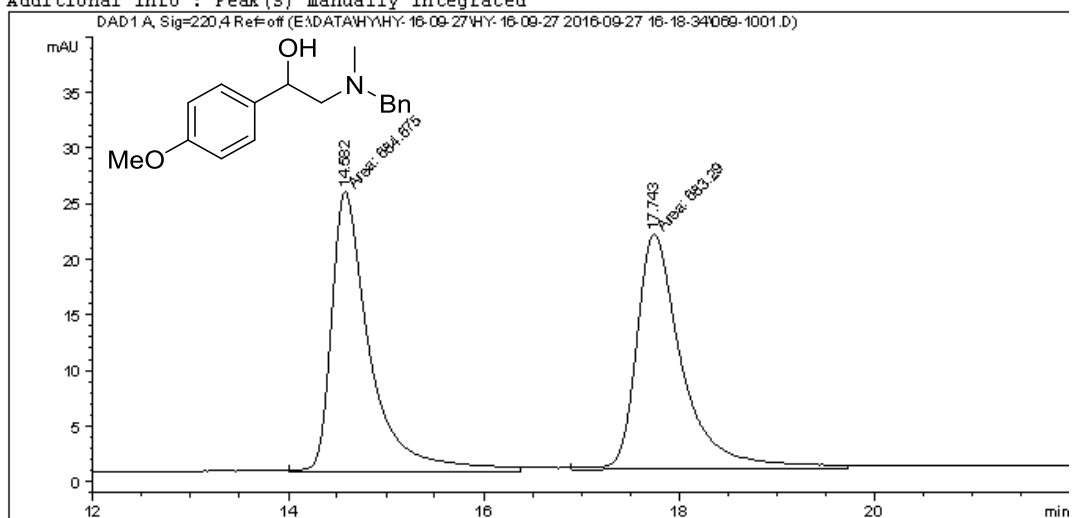
Totals : 1.13701e4 610.29456

=====
 *** End of Report ***

(S)-2-(Benzyl(methyl)amino)-1-(4-methoxyphenyl)ethanol (2f)

Data File E:\DATA\HY\HY-16-09-27\HY-16-09-27 2016-09-27 16-18-34\069-1001.D
Sample Name: HY-16-09-27-600-RAC

```
=====
Acq. Operator   : SYSTEM                      Seq. Line :   10
Acq. Instrument : 1260HPLC-DAD                Location  : Vial 69
Injection Date  : 9/27/2016 6:42:45 PM        Inj       :    1
                                           Inj Volume: 1.000 µl
Acq. Method     : E:\DATA\HY\HY-16-09-27\HY-16-09-27 2016-09-27 16-18-34\
DAD-OJ (1-6)-85-15
-1.0ML-ALLNM-15MIN.M
Last changed    : 9/27/2016 7:06:17 PM by SYSTEM
(modified after loading)
Analysis Method : E:\DATA\HY\HY-16-09-27\HY-16-09-27 2016-09-27 16-18-34\
DAD-OJ (1-6)-85-15
-1.0ML-ALLNM-15MIN.M (Sequence Method)
Last changed    : 2/9/2017 8:31:33 PM by SYSTEM
(modified after loading)
Additional Info : Peak(s) manually integrated
=====
```



Area Percent Report

```
Sorted By      : Signal
Multiplier     : 1.0000
Dilution       : 1.0000
Do not use Multiplier & Dilution Factor with ISTDs
```

Signal 1: DAD1 A, Sig=220,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	14.582	MM	0.4541	684.67493	25.13113	50.0506
2	17.743	MM	0.5397	683.29016	21.10250	49.9494

Totals : 1367.96509 46.23363

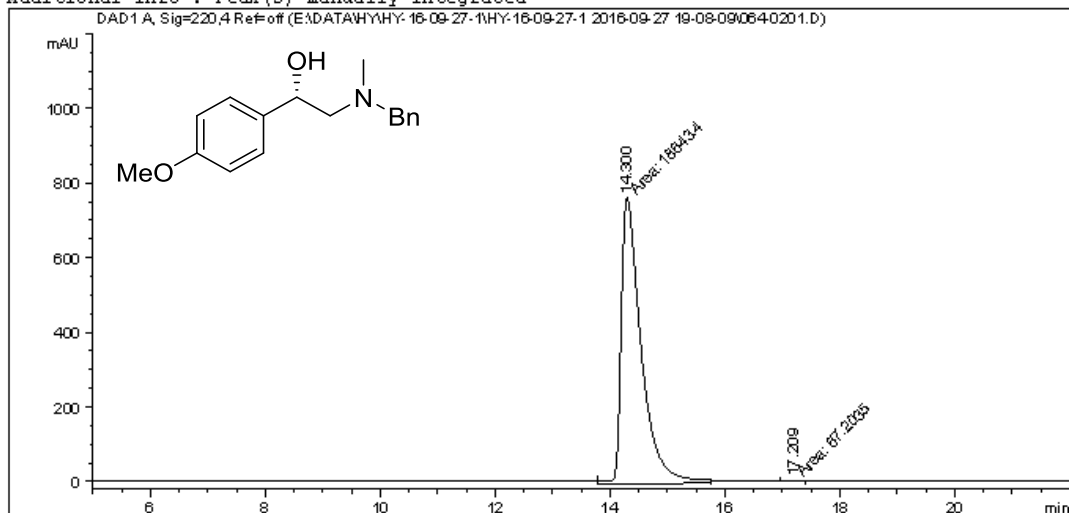
*** End of Report ***

```

=====
Acq. Operator   : SYSTEM                               Seq. Line :    2
Acq. Instrument : 1260HPLC-DAD                         Location  : Vial 64
Injection Date  : 9/27/2016 7:20:03 PM                Inj       :    1
                                                    Inj Volume: 1.000 µl

Acq. Method     : E:\DATA\HY\HY-16-09-27-1\HY-16-09-27-1 2016-09-27 19-08-09\DAD-0J(1-6)-
                  85-15-1.OML-ALLNM-15MIN.M
Last changed    : 9/27/2016 7:27:50 PM by SYSTEM
                  (modified after loading)
Analysis Method : E:\DATA\HY\HY-16-09-27-1\HY-16-09-27-1 2016-09-27 19-08-09\DAD-0J(1-6)-
                  85-15-1.OML-ALLNM-15MIN.M (Sequence Method)
Last changed    : 2/9/2017 8:37:13 PM by SYSTEM
                  (modified after loading)
  
```

Additional Info : Peak(s) manually integrated



=====
 Area Percent Report
 =====

```

Sorted By      :      Signal
Multiplier     :      1.0000
Dilution       :      1.0000
Do not use Multiplier & Dilution Factor with ISTDs
  
```

Signal 1: DAD1 A, Sig=220,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	14.300	MM	0.4055	1.86434e4	766.19769	99.6408
2	17.209	MM	0.3921	67.20346	2.85659	0.3592

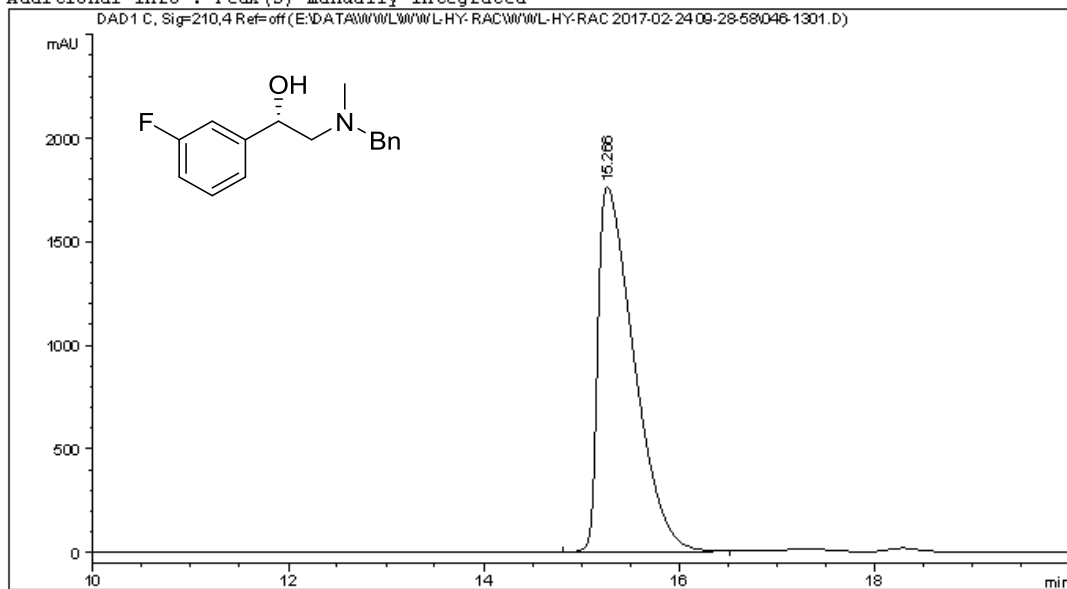
Totals : 1.87106e4 769.05429

=====
 *** End of Report ***

Data File E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\046-1301.D
Sample Name: M-F-EE

```
=====
Acq. Operator   : SYSTEM                               Seq. Line :   13
Acq. Instrument : 1260HPLC-DAD                         Location  : Vial 46
Injection Date  : 2/24/2017 3:02:15 PM                 Inj       :    1
                                                    Inj Volume: 2.000 µl

Acq. Method     : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\DAD-0J(1-6)-97-3-
                  LML-2UL-ALL-30MIN.M
Last changed    : 2/24/2017 9:45:54 AM by SYSTEM
Analysis Method : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\DAD-0J(1-6)-97-3-
                  LML-2UL-ALL-30MIN.M (Sequence Method)
Last changed    : 2/24/2017 4:18:55 PM by SYSTEM
                  (modified after loading)
Additional Info : Peak(s) manually integrated
=====
```



=====
Area Percent Report
=====

Sorted By : Signal
Multiplier : 1.0000
Dilution : 1.0000
Do not use Multiplier & Dilution Factor with ISTDs

Signal 1: DAD1 C, Sig=210,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	15.266	BB	0.3920	4.50202e4	1762.76770	100.0000

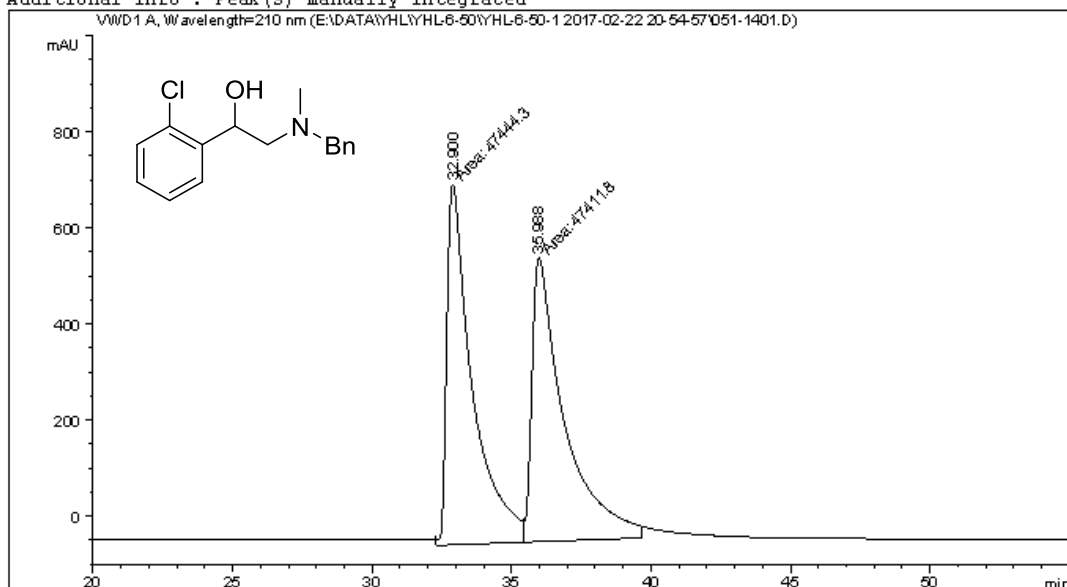
Totals : 4.50202e4 1762.76770

=====
*** End of Report ***

(S)-2-(Benzyl(methyl)amino)-1-(2-chlorophenyl)ethanol (2h)

Data File E:\DATA\YHL\YHL-6-50\YHL-6-50-1 2017-02-22 20-54-57\051-1401.D
Sample Name: HY-0-CL-RAC

```
=====
Acq. Operator   : SYSTEM                      Seq. Line :   14
Acq. Instrument : 1260HPLC-VWD                Location  : Vial 51
Injection Date  : 2/23/2017 6:30:51 AM       Inj       :    1
                                           Inj Volume: 1.000 µl
Acq. Method     : E:\DATA\YHL\YHL-6-50\YHL-6-50-1 2017-02-22 20-54-57\VWD-AD (1-2)-98-2-1UL
                                           -0.3ML-210NM-60MIN.M
Last changed    : 2/22/2017 9:42:42 PM by SYSTEM
Analysis Method : E:\DATA\YHL\YHL-6-50\YHL-6-50-1 2017-02-22 20-54-57\VWD-AD (1-2)-98-2-1UL
                                           -0.3ML-210NM-60MIN.M (Sequence Method)
Last changed    : 2/23/2017 8:13:44 PM by SYSTEM
                                           (modified after loading)
Additional Info  : Peak(s) manually integrated
=====
```



Area Percent Report

```
Sorted By      :      Signal
Multiplier     :      1.0000
Dilution       :      1.0000
Do not use Multiplier & Dilution Factor with ISTDs
```

Signal 1: VWD1 A, Wavelength=210 nm

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	32.900	MF	1.0537	4.74443e4	750.42676	50.0171
2	35.988	FM	1.3393	4.74118e4	589.99445	49.9829

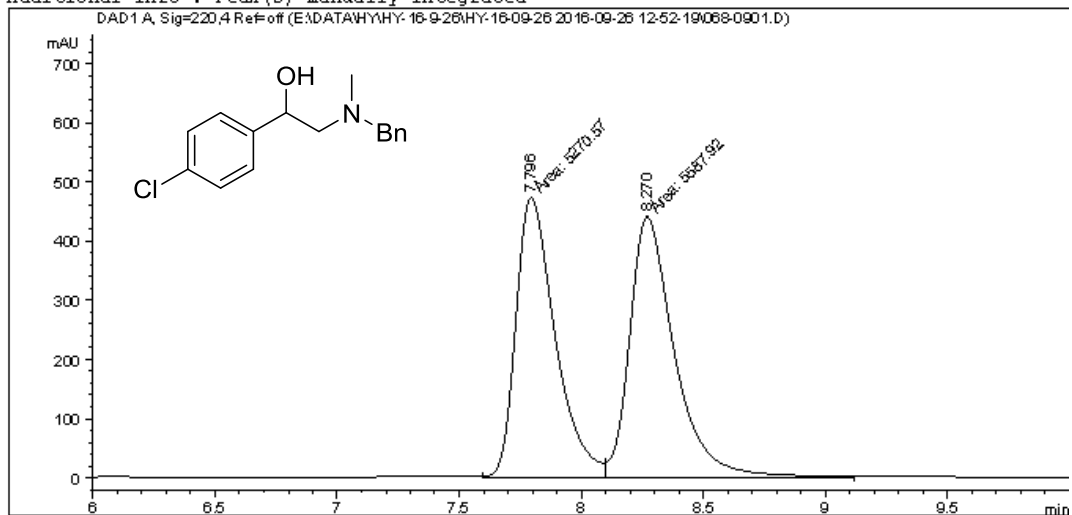
Totals : 9.48562e4 1340.42120

*** End of Report ***

(S)-2-(Benzyl(methyl)amino)-1-(4-chlorophenyl)ethanol (2i)

Data File E:\DATA\HY\HY-16-9-26\HY-16-09-26 2016-09-26 12-52-19\068-0901.D
Sample Name: HY-604

```
=====
Acq. Operator   : SYSTEM                      Seq. Line :    9
Acq. Instrument : 1260HPLC-DAD                Location  : Vial 68
Injection Date  : 9/26/2016 2:55:30 PM        Inj       :    1
                                           Inj Volume: 1.000 µl
Acq. Method     : E:\DATA\HY\HY-16-9-26\HY-16-09-26 2016-09-26 12-52-19\DAD-0J(1-6)-85-15-
                                           1.OML-ALLNM-15MIN.M
Last changed    : 9/26/2016 12:52:20 PM by SYSTEM
Analysis Method : E:\DATA\HY\HY-16-9-26\HY-16-09-26 2016-09-26 12-52-19\DAD-0J(1-6)-85-15-
                                           1.OML-ALLNM-15MIN.M (Sequence Method)
Last changed    : 2/9/2017 9:04:21 PM by SYSTEM
                                           (modified after loading)
Additional Info  : Peak(s) manually integrated
=====
```



Area Percent Report

```
Sorted By      :      Signal
Multiplier     :      1.0000
Dilution       :      1.0000
Do not use Multiplier & Dilution Factor with ISTDs
```

Signal 1: DAD1 A, Sig=220,4 Ref=off

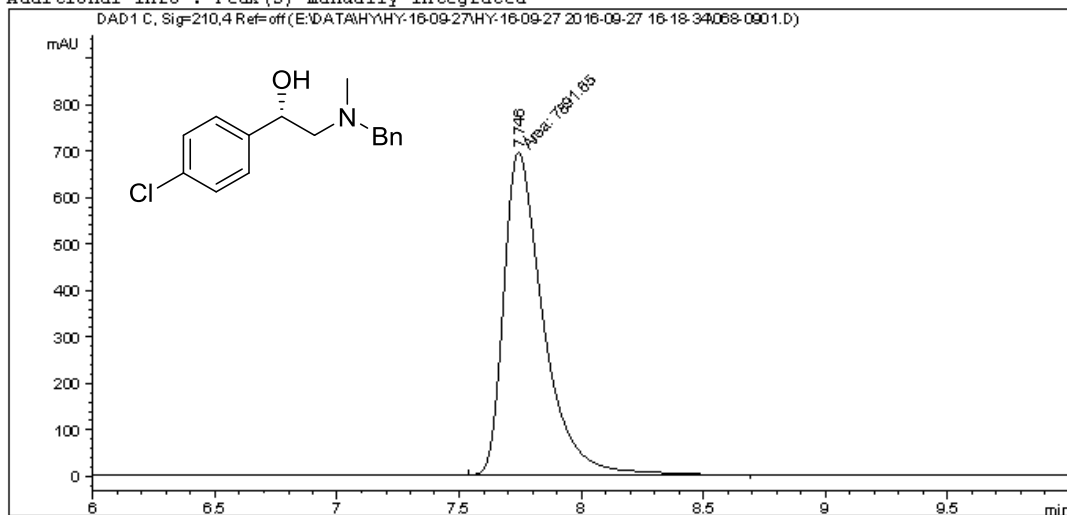
Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	7.96	MF	0.1856	5270.57324	473.17984	48.5387
2	8.270	FM	0.2114	5587.92285	440.55948	51.4613

Totals : 1.08585e4 913.73932

*** End of Report ***

Data File E:\DATA\HY\HY-16-09-27\HY-16-09-27 2016-09-27 16-18-34\068-0901.D
Sample Name: HY-16-09-27-604

```
=====
Acq. Operator   : SYSTEM                               Seq. Line :    9
Acq. Instrument : 1260HPLC-DAD                         Location  : Vial 68
Injection Date  : 9/27/2016 6:26:51 PM                 Inj       :    1
                                                    Inj Volume: 1.000 µl
Acq. Method     : E:\DATA\HY\HY-16-09-27\HY-16-09-27 2016-09-27 16-18-34\DAD-OJ(1-6)-85-15
                  -1.0ML-ALLNM-15MIN.M
Last changed    : 9/27/2016 4:18:35 PM by SYSTEM
Analysis Method : E:\DATA\HY\HY-16-09-27\HY-16-09-27 2016-09-27 16-18-34\DAD-OJ(1-6)-85-15
                  -1.0ML-ALLNM-15MIN.M (Sequence Method)
Last changed    : 2/9/2017 9:15:51 PM by SYSTEM
                  (modified after loading)
Additional Info : Peak(s) manually integrated
=====
```



=====
Area Percent Report
=====

Sorted By : Signal
Multiplier : 1.0000
Dilution : 1.0000
Do not use Multiplier & Dilution Factor with ISTDs

Signal 1: DAD1 C, Sig=210,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	7.746	MM	0.1894	7891.65234	694.37036	100.0000

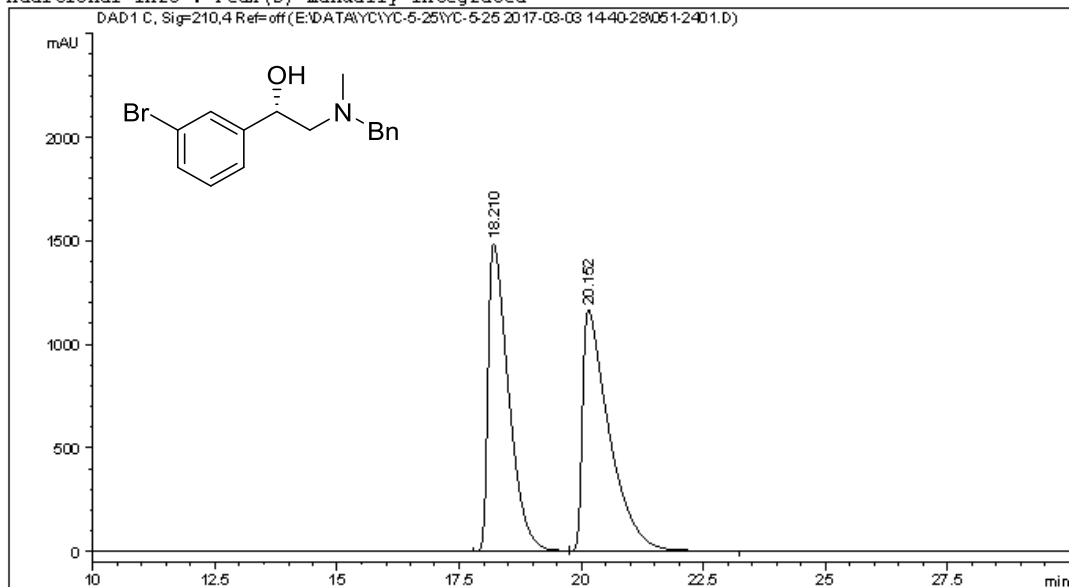
Totals : 7891.65234 694.37036

=====
*** End of Report ***

(S)-2-(Benzyl(methyl)amino)-1-(3-bromophenyl)ethanol (2j)

Data File E:\DATA\YC\YC-5-25\YC-5-25 2017-03-03 14-40-28\051-2401.D
Sample Name: HY-M-BR-RAC

```
=====
Acq. Operator   : SYSTEM                      Seq. Line :   24
Acq. Instrument : 1260HPLC-DAD                Location  : Vial 51
Injection Date  : 3/4/2017 5:21:00 AM        Inj       :    1
                                           Inj Volume: 2.000 µl
Acq. Method     : E:\DATA\YC\YC-5-25\YC-5-25 2017-03-03 14-40-28\DAD-0J(1-6)-97-3-1ML-2UL-
                  ALL-45MIN.M
Last changed    : 3/3/2017 8:47:17 PM by SYSTEM
Analysis Method : E:\DATA\YC\YC-5-25\YC-5-25 2017-03-03 14-40-28\DAD-0J(1-6)-97-3-1ML-2UL-
                  ALL-45MIN.M (Sequence Method)
Last changed    : 3/11/2017 9:40:01 AM by SYSTEM
                  (modified after loading)
Additional Info : Peak(s) manually integrated
=====
```



Area Percent Report

```
Sorted By      :      Signal
Multiplier     :      1.0000
Dilution       :      1.0000
Do not use Multiplier & Dilution Factor with ISTDs
```

Signal 1: DAD1 C, Sig=210,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	18.210	VV	0.4345	4.33977e4	1486.92542	49.4453
2	20.152	VB	0.5425	4.43713e4	1164.65344	50.5547

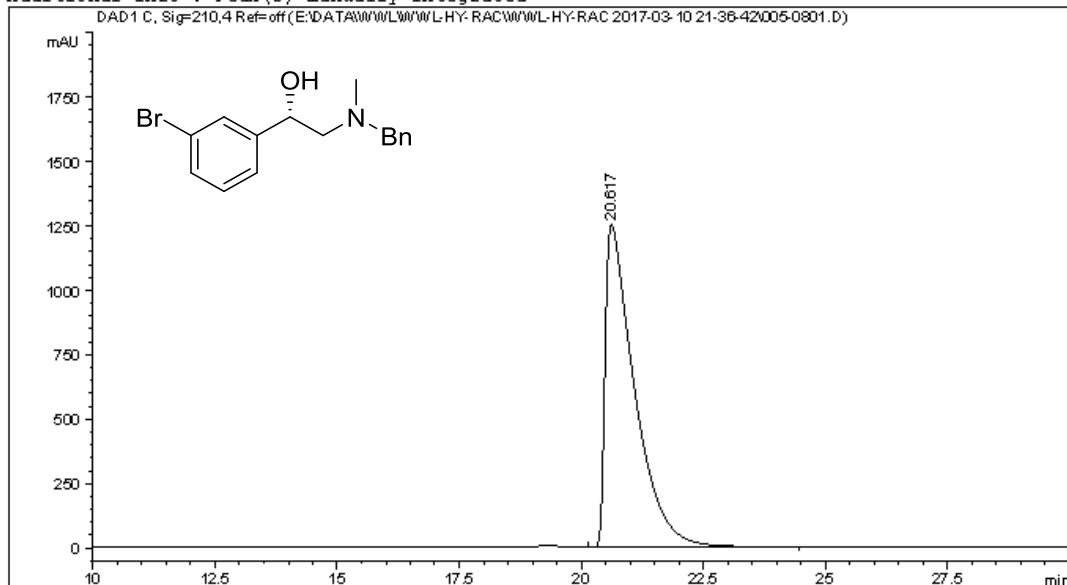
Totals : 8.77690e4 2651.57886

*** End of Report ***

Data File E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-03-10 21-36-42\005-0801.D
Sample Name: WVL-HY-M-BR-EE

```
=====
Acq. Operator   : SYSTEM                      Seq. Line :    8
Acq. Instrument : 1260HPLC-DAD                Location  : Vial 5
Injection Date  : 3/11/2017 12:07:34 AM      Inj       :    1
                                                Inj Volume: 2.000 µl

Acq. Method     : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-03-10 21-36-42\DAD-0J(1-6)-97-3-
                  LML-2UL-ALL-45MIN.M
Last changed    : 3/10/2017 10:26:20 PM by SYSTEM
Analysis Method : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-03-10 21-36-42\DAD-0J(1-6)-97-3-
                  LML-2UL-ALL-45MIN.M (Sequence Method)
Last changed    : 3/11/2017 9:44:35 AM by SYSTEM
                  (modified after loading)
Additional Info : Peak(s) manually integrated
=====
```



=====
Area Percent Report
=====

Sorted By : Signal
Multiplier : 1.0000
Dilution : 1.0000
Do not use Multiplier & Dilution Factor with ISTDs

Signal 1: DAD1 C, Sig=210,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	20.617	BB	0.5934	5.22552e4	1250.82336	100.0000

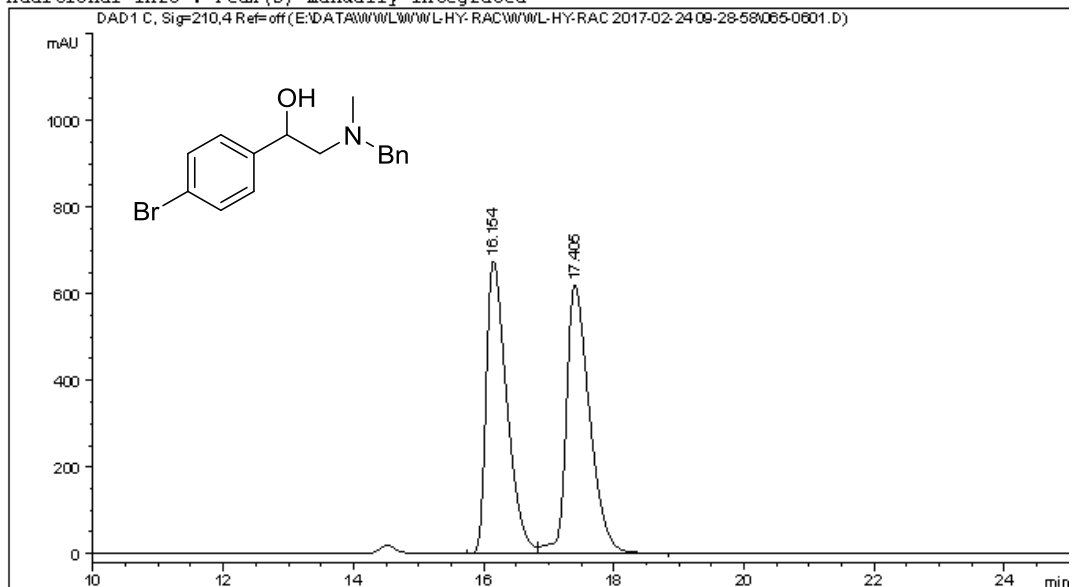
Totals : 5.22552e4 1250.82336

=====
*** End of Report ***

(S)-2-(Benzyl(methyl)amino)-1-(4-bromophenyl)ethanol (2k)

Data File E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\065-0601.D
Sample Name: P-BR-RAC

```
=====
Acq. Operator   : SYSTEM                      Seq. Line :    6
Acq. Instrument : 1260HPLC-DAD                Location  : Vial 65
Injection Date  : 2/24/2017 11:45:43 AM      Inj       :    1
                                           Inj Volume: 2.000 µl
Acq. Method     : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\DAD-0J(1-6)-95-5-
                  LML-2UL-ALL-30MIN.M
Last changed    : 2/24/2017 9:28:58 AM by SYSTEM
Analysis Method : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\DAD-0J(1-6)-95-5-
                  LML-2UL-ALL-30MIN.M (Sequence Method)
Last changed    : 2/24/2017 4:30:39 PM by SYSTEM
                  (modified after loading)
Additional Info  : Peak(s) manually integrated
=====
```



Area Percent Report

```
Sorted By      :      Signal
Multiplier     :      1.0000
Dilution       :      1.0000
Do not use Multiplier & Dilution Factor with ISTDs
```

Signal 1: DAD1 C, Sig=210,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	16.154	BV	0.3347	1.48655e4	675.62274	49.1446
2	17.405	VB	0.3757	1.53830e4	619.44073	50.8554

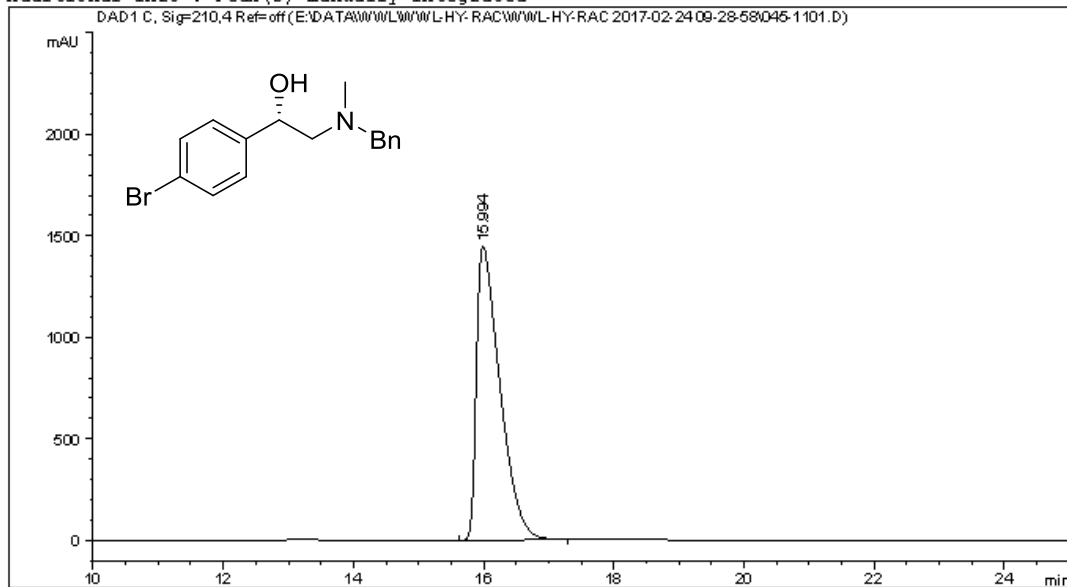
Totals : 3.02485e4 1295.06348

*** End of Report ***

```

=====
Acq. Operator   : SYSTEM                      Seq. Line : 11
Acq. Instrument : 1260HPLC-DAD                Location  : Vial 45
Injection Date  : 2/24/2017 2:20:14 PM        Inj       : 1
                                           Inj Volume: 2.000 µl

Acq. Method     : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\DAD-0J(1-6)-95-5-
                  1ML-2UL-ALL-30MIN.M
Last changed    : 2/24/2017 9:28:58 AM by SYSTEM
Analysis Method : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-24 09-28-58\DAD-0J(1-6)-95-5-
                  1ML-2UL-ALL-30MIN.M (Sequence Method)
Last changed    : 2/24/2017 4:33:15 PM by SYSTEM
                  (modified after loading)
Additional Info : Peak(s) manually integrated
  
```



=====
 Area Percent Report
 =====

```

Sorted By      : Signal
Multiplier     : 1.0000
Dilution       : 1.0000
Do not use Multiplier & Dilution Factor with ISTDs
  
```

Signal 1: DAD1 C, Sig=210,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	15.994	BB	0.3777	3.61847e4	1447.10156	100.0000

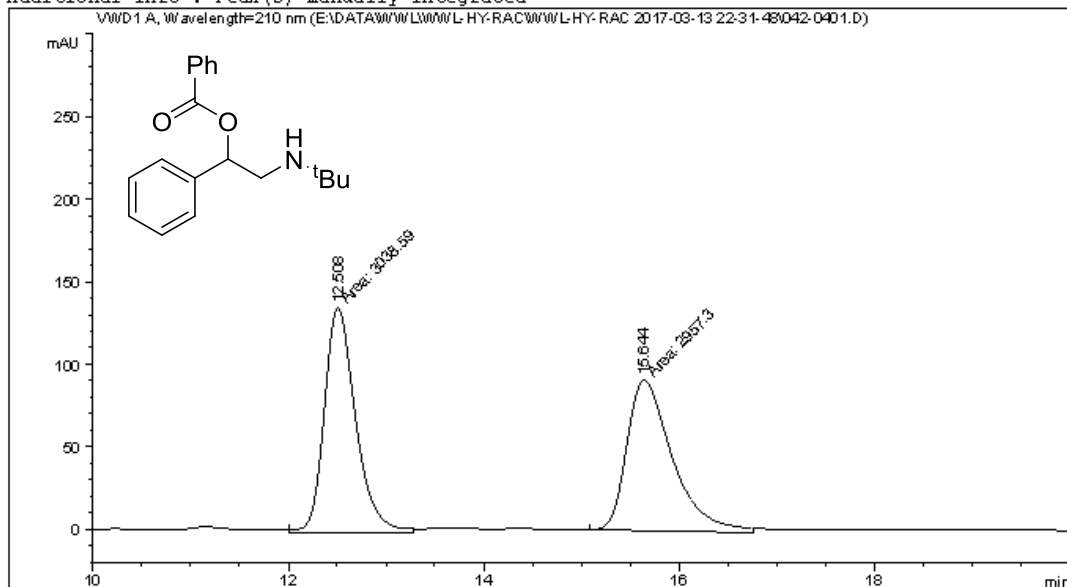
Totals : 3.61847e4 1447.10156

=====
 *** End of Report ***

(S)-2-(tert-butylamino)-1-phenylethanol (2l)

Data File E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-03-13 22-31-48\042-0401.D
Sample Name: WWK-HY-TBU-RAC-1

```
=====
Acq. Operator   : SYSTEM                      Seq. Line :    4
Acq. Instrument : 1260HPLC-VWD                Location  : Vial 42
Injection Date  : 3/13/2017 11:34:54 PM      Inj       :    1
                                           Inj Volume: 1.000 µl
Acq. Method     : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-03-13 22-31-48\VWD-AD(1-2)-85-15-
                  LML-1UL-210NM-30MIN.M
Last changed    : 3/13/2017 10:31:49 PM by SYSTEM
Analysis Method : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-03-13 22-31-48\VWD-AD(1-2)-85-15-
                  LML-1UL-210NM-30MIN.M (Sequence Method)
Last changed    : 3/14/2017 4:30:47 PM by SYSTEM
                  (modified after loading)
Additional Info : Peak(s) manually integrated
=====
```



Area Percent Report

```
Sorted By      : Signal
Multiplier     : 1.0000
Dilution       : 1.0000
Do not use Multiplier & Dilution Factor with ISTDs
```

Signal 1: VWD1 A, Wavelength=210 nm

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	12.508	MM	0.3712	3038.59009	136.44746	50.6779
2	15.644	MM	0.5403	2957.30371	91.22073	49.3221

Totals : 5995.89380 227.66819

*** End of Report ***

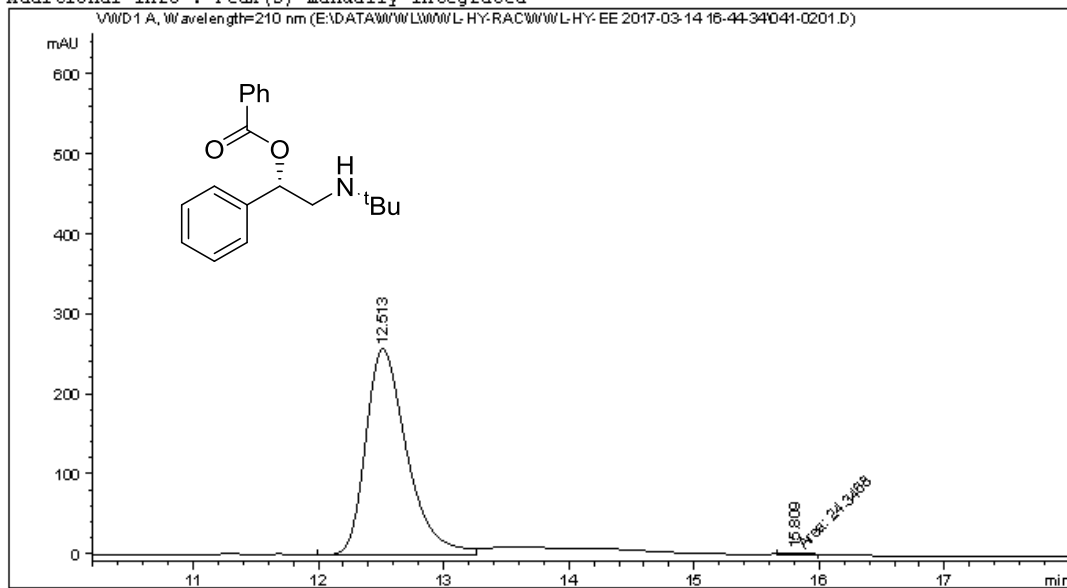
Data File E:\DATA\WVL\WVL-HY-RAC\WVL-HY-EE 2017-03-14 16-44-34\041-0201.D
 Sample Name: WVL-HY-TBU-EE

```

=====
Acq. Operator   : SYSTEM                               Seq. Line :    2
Acq. Instrument : 1260HPLC-VWD                         Location  : Vial 41
Injection Date  : 3/14/2017 4:56:09 PM                Inj       :    1
                                                    Inj Volume: 5.000 µl

Acq. Method     : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-EE 2017-03-14 16-44-34\WVD-ADH(1-2)-85-15-
                  1-SUL-210NM-30MIN.M
Last changed    : 3/14/2017 5:23:07 PM by SYSTEM
                  (modified after loading)
Analysis Method : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-EE 2017-03-14 16-44-34\WVD-ADH(1-2)-85-15-
                  1-SUL-210NM-30MIN.M (Sequence Method)
Last changed    : 3/14/2017 5:28:57 PM by SYSTEM
                  (modified after loading)
  
```

Additional Info : Peak(s) manually integrated



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 Area Percent Report
 =====

```

Sorted By      :      Signal
Multiplier     :      1.0000
Dilution      :      1.0000
Do not use Multiplier & Dilution Factor with ISTDs
  
```

Signal 1: WVD1 A, Wavelength=210 nm

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	12.513	BV	0.3473	5871.37012	257.20615	99.5870
2	15.809	MM	0.2398	24.34677	1.69225	0.4130

Totals : 5895.71689 258.89840

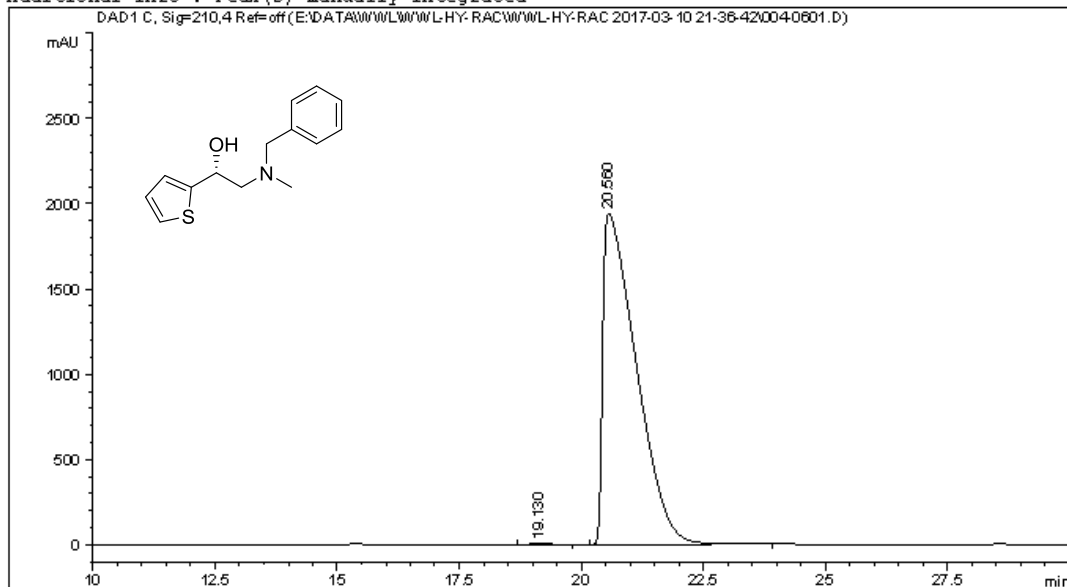
Data File E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-03-10 21-36-42\004-0601.D
 Sample Name: WVL-HY-SAIFEN-EE

```

=====
Acq. Operator   : SYSTEM                      Seq. Line :    6
Acq. Instrument : 1260HPLC-DAD                Location  : Vial 4
Injection Date  : 3/10/2017 11:25:36 PM      Inj       :    1
                                           Inj Volume: 2.000 µl

Acq. Method     : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-03-10 21-36-42\DAD-0J(1-6)-95-5-
                  LML-2UL-ALL-30MIN.M
Last changed    : 3/10/2017 10:24:42 PM by SYSTEM
Analysis Method : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-03-10 21-36-42\DAD-0J(1-6)-95-5-
                  LML-2UL-ALL-30MIN.M (Sequence Method)
Last changed    : 3/11/2017 9:35:39 AM by SYSTEM
                  (modified after loading)
  
```

Additional Info : Peak(s) manually integrated



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 Area Percent Report
 =====

```

Sorted By      :      Signal
Multiplier     :      1.0000
Dilution       :      1.0000
Do not use Multiplier & Dilution Factor with ISTDs
  
```

Signal 1: DAD1 C, Sig=210,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	19.130	BB	0.3338	110.10705	4.72548	0.1197
2	20.560	BB	0.5945	9.18732e4	1939.86816	99.8803

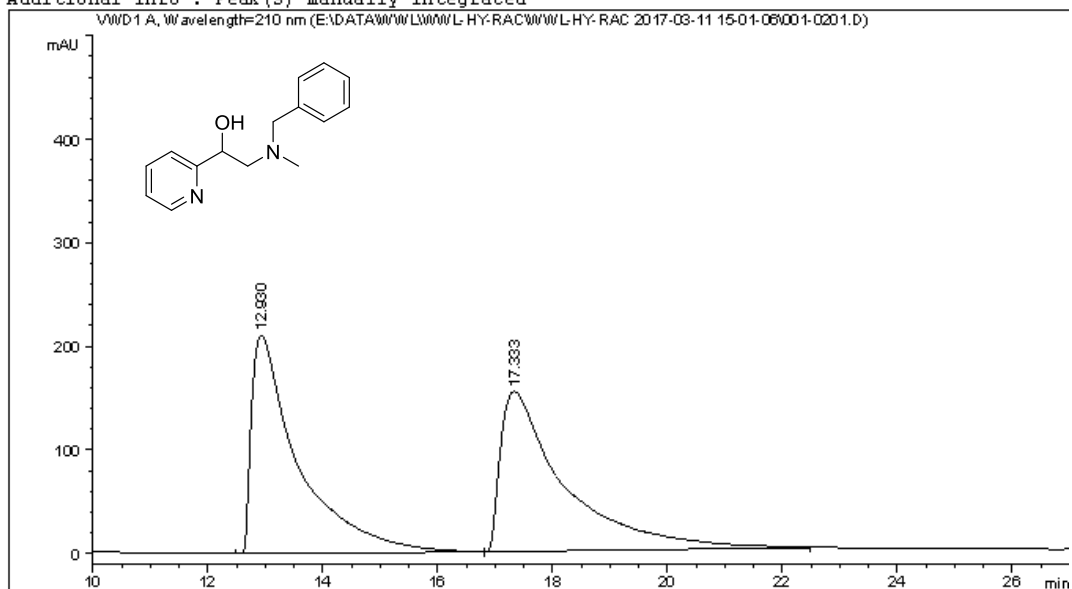
Totals : 9.19833e4 1944.59365

=====
 *** End of Report ***

(R)-2-(benzyl(methyl)amino)-1-(pyridin-2-yl)ethanol (2n)

Data File E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-03-11 15-01-06\001-0201.D
Sample Name: WVL-HY-PY-RAC

```
=====
Acq. Operator   : SYSTEM                      Seq. Line :    2
Acq. Instrument : 1260HPLC-VWD                Location  : Vial 1
Injection Date  : 3/11/2017 3:12:39 PM        Inj       :    1
                                           Inj Volume: 1.000 µl
Acq. Method     : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-03-11 15-01-06\VWD-AS(1-6)-97-3-
                  LML-1UL-210NM-40MIN.M
Last changed    : 3/11/2017 3:01:06 PM by SYSTEM
Analysis Method : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-03-11 15-01-06\VWD-AS(1-6)-97-3-
                  LML-1UL-210NM-40MIN.M (Sequence Method)
Last changed    : 3/11/2017 6:40:28 PM by SYSTEM
                  (modified after loading)
Additional Info : Peak(s) manually integrated
=====
```



Area Percent Report

```
Sorted By      : Signal
Multiplier     : 1.0000
Dilution       : 1.0000
Do not use Multiplier & Dilution Factor with ISTDs
```

Signal 1: VWD1 A, Wavelength=210 nm

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	12.930	BB	0.7926	1.19120e4	211.00032	49.8365
2	17.333	BB	1.0925	1.19901e4	154.47755	50.1635

Totals : 2.39021e4 365.47787

*** End of Report ***

(S)-3-(2-(benzyl(methyl)amino)-1-hydroxyethyl)phenol (2o)

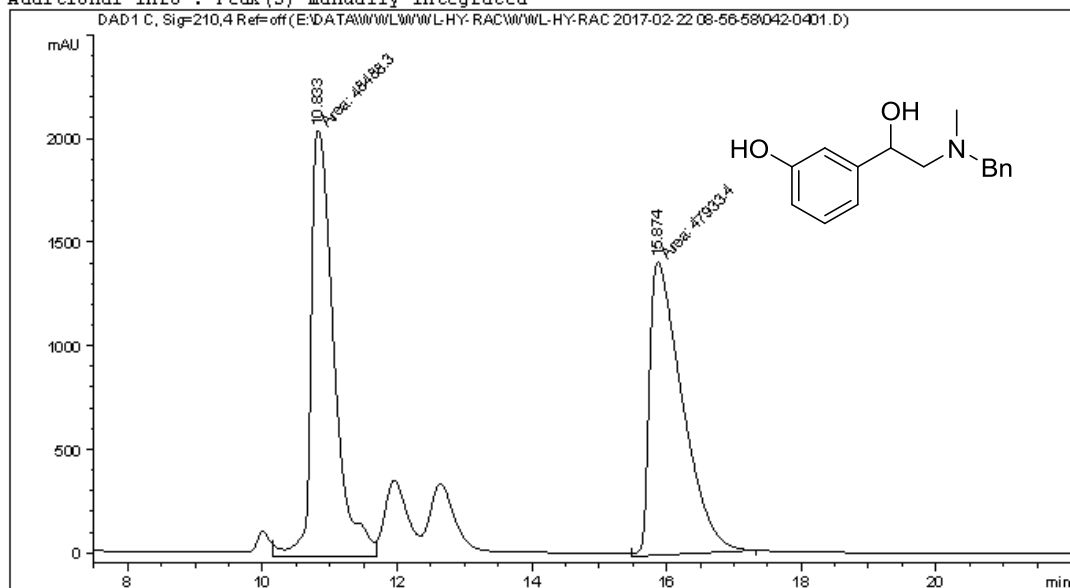
Data File E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-22 08-56-58\042-0401.D
Sample Name: HY-M-OH-RAC

=====

Acq. Operator	: SYSTEM	Seq. Line	: 4
Acq. Instrument	: 1260HPLC-DAD	Location	: Vial 42
Injection Date	: 2/22/2017 9:57:24 AM	Inj	: 1
		Inj Volume	: 2.000 µl

Acq. Method : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-22 08-56-58\DAD-0J(1-6)-80-20-1ML-2UL-ALL-30MIN.M
Last changed : 2/22/2017 10:24:57 AM by SYSTEM
(modified after loading)
Analysis Method : E:\DATA\WVL\WVL-HY-RAC\WVL-HY-RAC 2017-02-22 08-56-58\DAD-0J(1-6)-80-20-1ML-2UL-ALL-30MIN.M (Sequence Method)
Last changed : 2/24/2017 10:00:38 AM by SYSTEM
(modified after loading)

Additional Info : Peak(s) manually integrated



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Area Percent Report
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Sorted By : Signal
Multiplier : 1.0000
Dilution : 1.0000
Do not use Multiplier & Dilution Factor with ISTDs

Signal 1: DAD1 C, Sig=210,4 Ref=off

Peak #	RetTime [min]	Type	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	10.833	MM	0.3931	4.84883e4	2055.69678	50.2878
2	15.874	MM	0.5646	4.79334e4	1415.00256	49.7122

Totals : 9.64216e4 3470.69934

7. Reference

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