

Synthesis of a New Chiral Cyclic β -Amino Alcohol

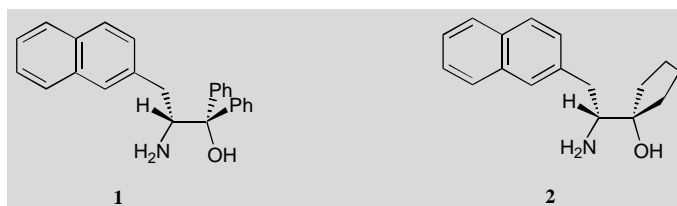
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Abstract: A new chiral cyclic β -amino alcohol (*R*)-1-(1'-amino-2'-naphthylethyl)-cyclopentanol **2**, which had not been reported previously, was prepared from (*R*)-3-(2-naphthyl)-alanine **3**.

Keywords: β -Amino alcohol, chiral, cyclic.

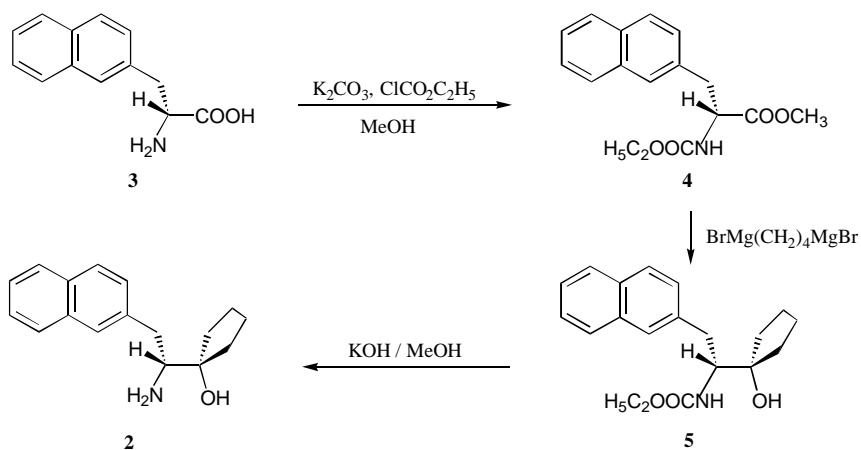
The enantioselective chiral 1,3,2-oxazaborolidine-catalyzed reduction of prochiral ketones by borane has been intensively investigated¹⁻³. Most oxazaborolidine catalysts have been prepared from chiral β -amino alcohols. In our previous study on the catalyst structure-enantioselectivity relationships, we prepared (*R*)-2-amino-1,1-diphenyl-3-(2-naphthyl)-1-propanol **1** which proved to be a good precursor of chiral catalyst for the enantioselective borane reduction of prochiral ketones⁴⁻⁹. This letter describes the synthesis of a new chiral cyclic β -amino alcohol (*R*)-1-(1'-amino-2'-naphthylethyl)-cyclopentanol **2** from (*R*)-3-(2-naphthyl)-alanine **3**. According to Marttens¹⁰, the less flexible **2** may achieve more efficient enantioselectivity than **1** does. The synthesis of **2** is shown in **Scheme 1**.



The reaction of **3** with ethyl chloroformate in MeOH with K₂CO₃ gave the ethyl ester of the *N*-ethoxycarbonyl amino acid **4** in one-pot reaction¹¹. Treatment of **4** with excess Grignard reagent prepared from 1,4-dibromobutane afforded *N*-ethoxycarbonyl cyclic β -amino alcohol **5**, which, after deprotection by alkaline hydrolysis gave the target compound **2**, m.p. 64–66°C, [α]_D²⁰ 9.97(CHCl₃, c = 1.72). Elemental anal. calcd for C₁₇H₂₁NO: C, 79.96; H, 8.29; N, 5.49. Found: C, 79.65; H, 8.48; N, 5.50. ¹H NMR (δ ppm, CDCl₃): 1.45 (s, 2H, NH₂), 1.70 (m, 8H, CH₂CH₂CH₂CH₂), 1.92 (s, 1H, OH), 2.66 (m, 1H, NCH), 3.10 (m, 2H, ArCH₂), 7.22–7.80 (m, 7H, ArH).

Further investigation of the enantioselective reduction of prochiral ketones catalyzed by oxazaborolidine formed by **2** would be reported soon.

Scheme 1



Acknowledgment

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