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Doctoral dissertation: "Synthesis and transformations of chiral β-amino alcohols with pyridine and 1-phenylethylamine fragments."

Abstract

The project was aimed at the design and development of novel modular chiral ligandscatalysts for asymmetric reactions. The key concept was an introduction of additional metal complexing moieties of pyridine and 2,2'-bipyridine to the chiral scaffold offered by α phenylethylamine (α -PEA). We synthesized new pyridine-substituted epoxides that were subsequently opened with α -PEA in the Sc(TfO)₃ – catalyzed reaction. The prepared diastereometric β -amino alcohols were successfully separated. In the next step, these products were transformed into the respective chiral diamines via azides. In the case of an α -pyridine amino alcohol, the first formed product was the aziridine derivative, then stereoselectively opened to the azido-amine. The Staudinger reduction of azides gave the desired products. The chiral diamines reacting with the appropriate salicylaldehydes were converted into the corresponding mono-aldimines. We also synthesized sulfur- and selenium-containing chiral ligands with phenylsulfanyl (-SPh) and phenylselenyl (-SePh) groups. The other experiments have shown that the corresponding cyclic sulfin- and sulfonamides could be obtained from chiral β -amino alcohols by simple reaction with thionyl chloride. These cyclic synthetic intermediates underwent the stereospecific S_N2 reactions, and the desired substitution products were formed. The obtained modular catalysts were tested in several asymmetric reactions: aldol, Henry, Michael reactions, sulfoxidation, and Tsuji-Trost reactions. The last one, the Pdcatalyzed process, led to up to 75% ee. The observed sense of asymmetric induction resulted from the nucleophilic attack at the position trans vs. the coordinated pyridine nitrogen.