

Natural Product Synthesis

Total Synthesis of the Indole Alkaloid (±)-and (+)- Methyl *N*- Decarbomethoxychanofruticosinate

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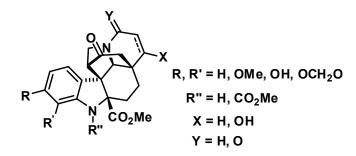




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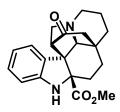
Introduction

- Methyl chanofruticosinates belong to growing family of indole alkaloids
- * Theses alkaloids have been isolated from Kopsia species (Apocynaceae)

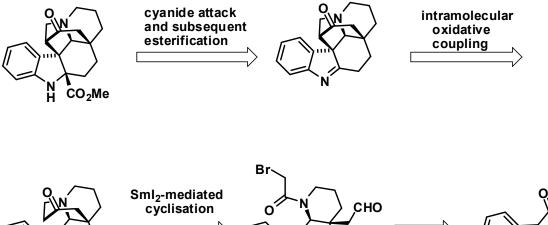




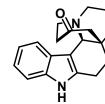
- widely distributed in tropical Asia and they have been historically used for the treatment of rheumatoid arthritis
- In particular, Methyl N- decarbomethoxychanofruticosinte displays antitussive activity in a citric acid induced guinea pig cough model and also shows relaxation activity against phenylephrine-induced contractions of rat aortas



Retrosynthetic analysis

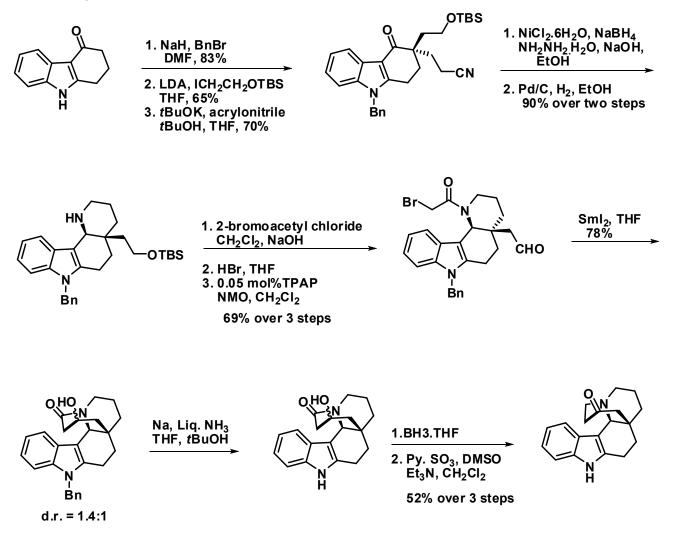


N R N

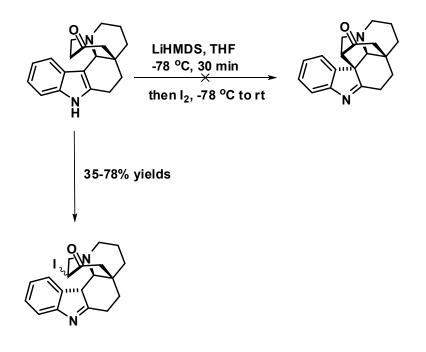


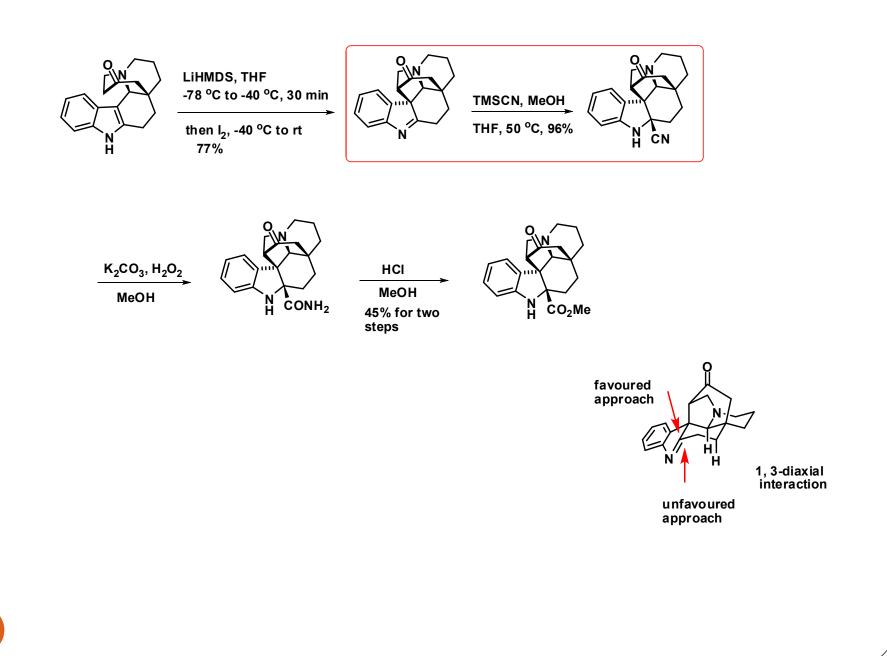
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Racemic Synthesis

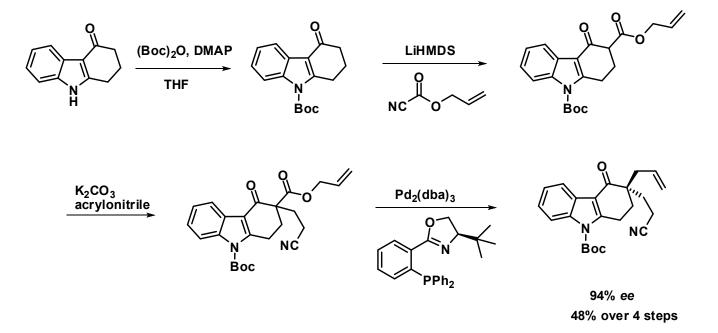


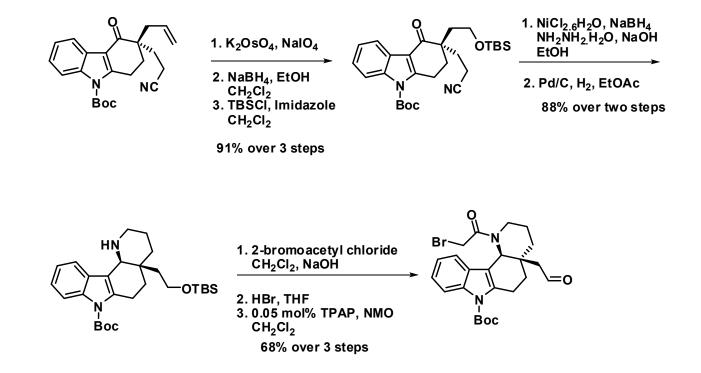
Iodination is much faster than Oxidation of enolate?

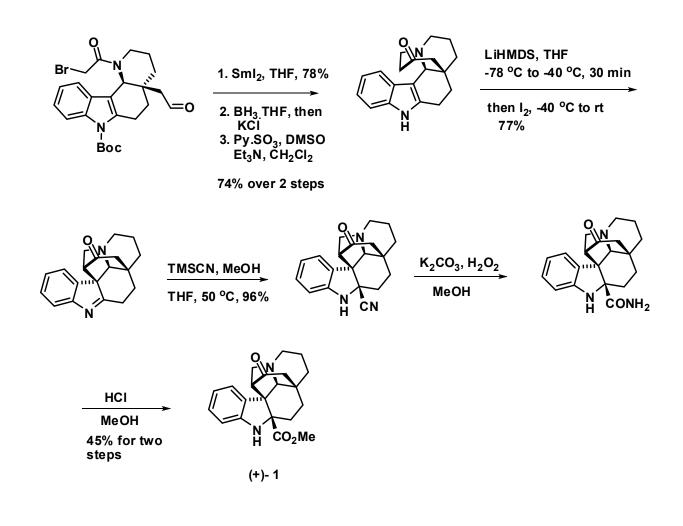




Preparation Starting material for enantioselective Synthesis







Conclusion and key features

- achieved the total synthesis of Methyl N- Decarbomethoxychanofruticosinate in both racemic and enantioselective forms for the first time
- ✤ racemic synthesis- 16 linear steps and enantioselective synthesis- 19 linear steps
- SmI2-mediated intramolecular Reformatsky-like reaction to create seven- membered ring and Intramolecular oxidative coupling to install caged and strained ring system

Thank you For your attention

