

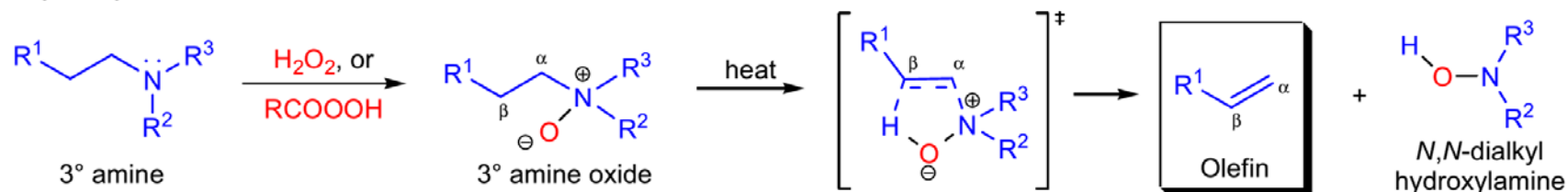
COPE ELIMINATION (COPE REACTION)

VS Hofmann elimination

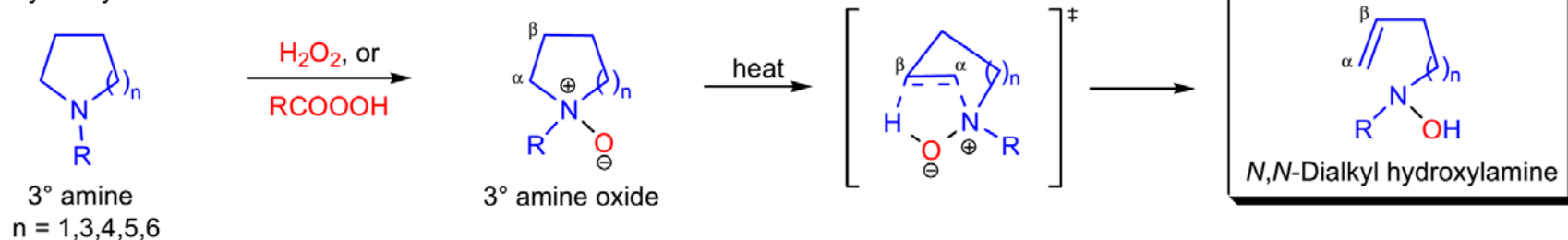
Xiaojun Han
2014-09-23

In 1949, A.C. Cope and co-workers discovered that by heating trialkylamine-N-oxides having hydrogens in the β -position, an olefin and N,N-dialkylhydroxylamine are formed. The transformation involving the stereoselective syn elimination of tertiary amine oxides is now referred to as the Cope elimination or Cope reaction.

Acyclic systems:

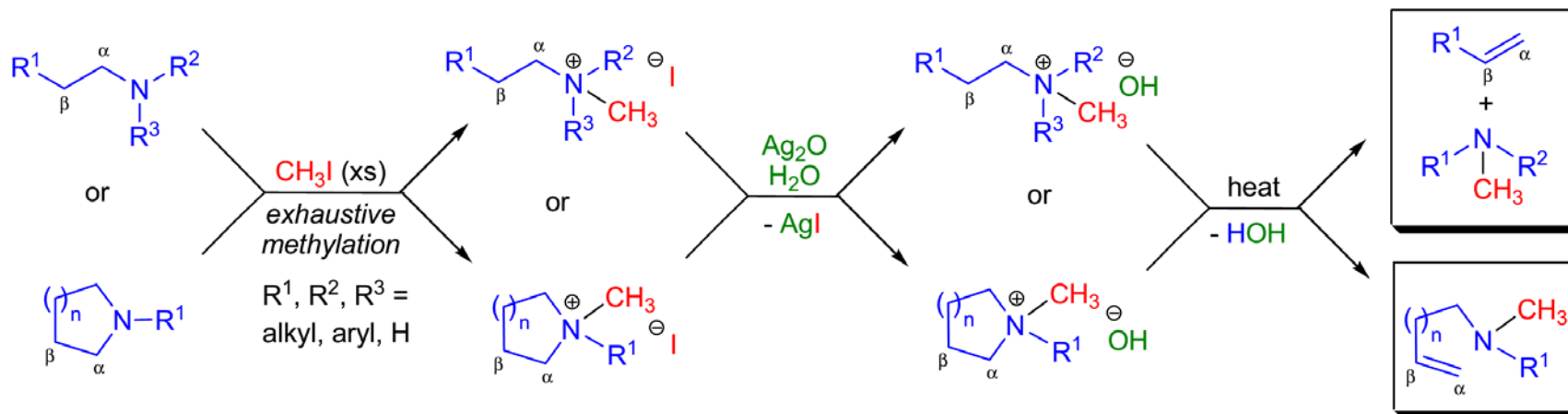


Cyclic systems:

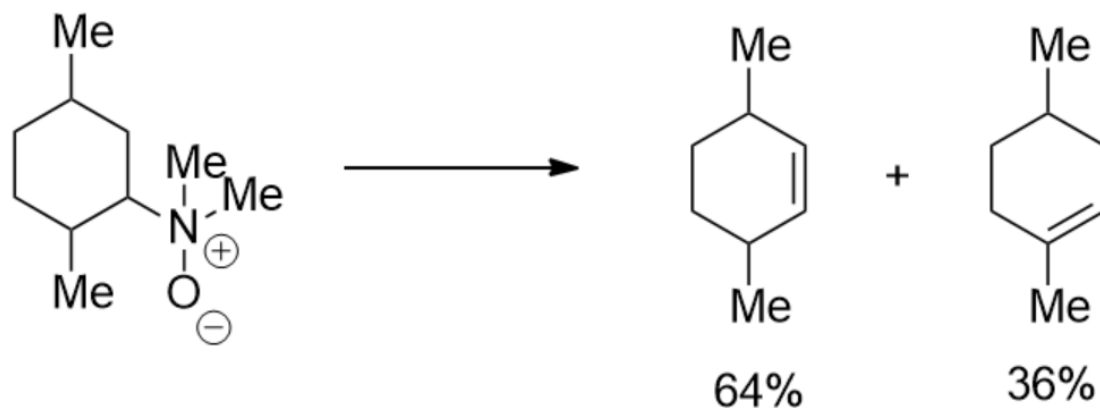
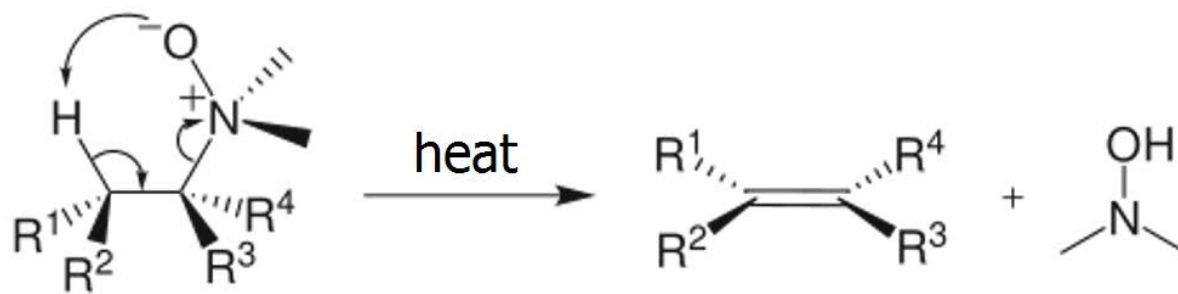


Hofmann elimination

The pyrolytic degradation of quaternary ammonium hydroxides to give a tertiary amine, an olefin and water is known as the Hofmann elimination.



Piperidines are resistant to an intramolecular Cope reaction but with pyrrolidine and with rings of size 7 and larger, the reaction product is an unsaturated hydroxyl amine. This result is consistent with the 5-membered cyclic transition state.



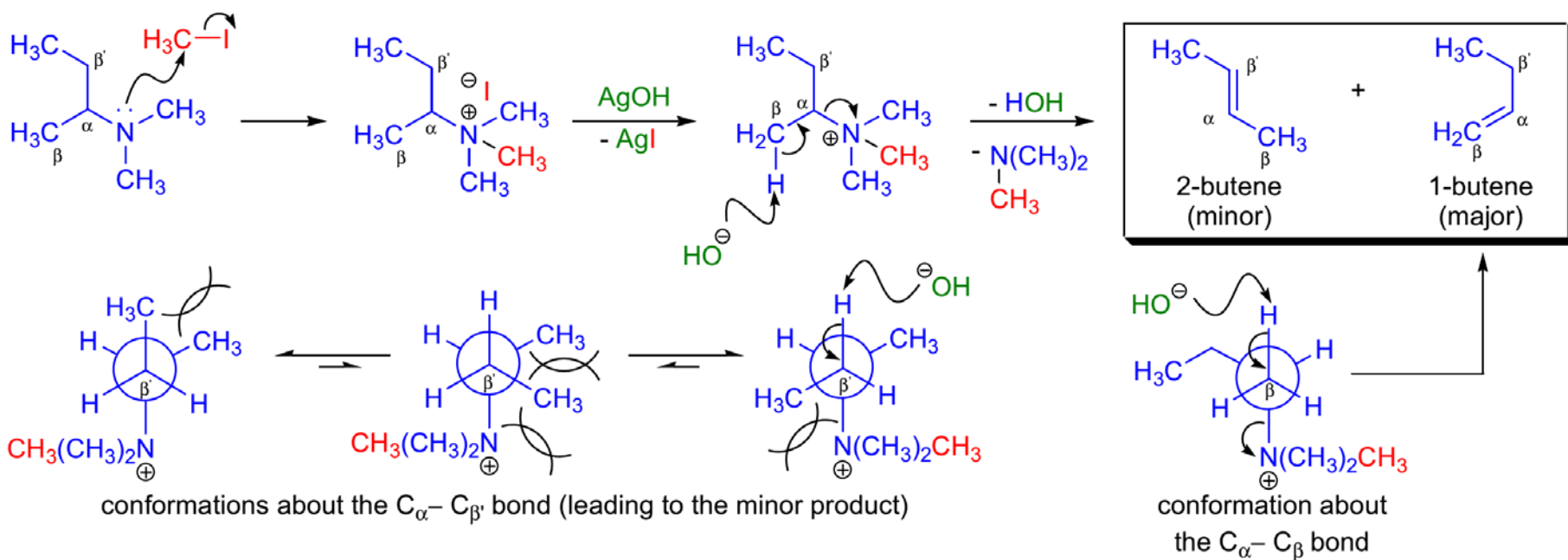
Mechanism (Cope elimination):

The reaction mechanism involves an intramolecular 5-membered cyclic transition state, leading to a syn elimination product.

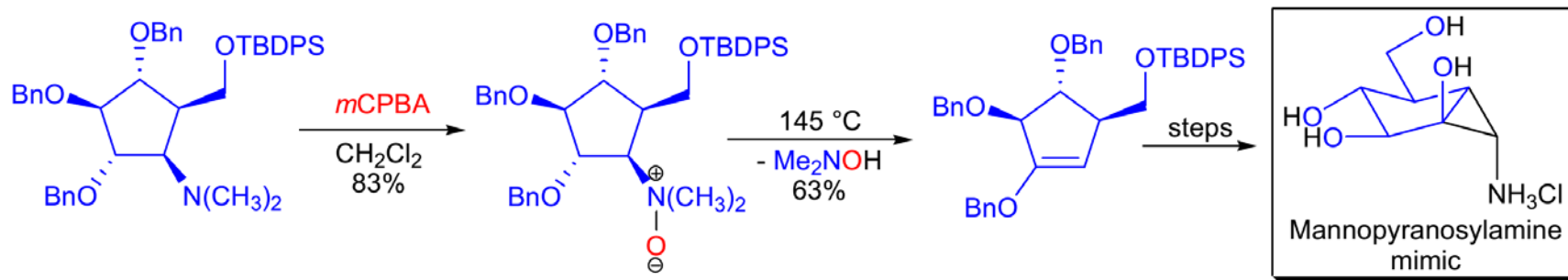


Mechanism (Hofmann elimination) :

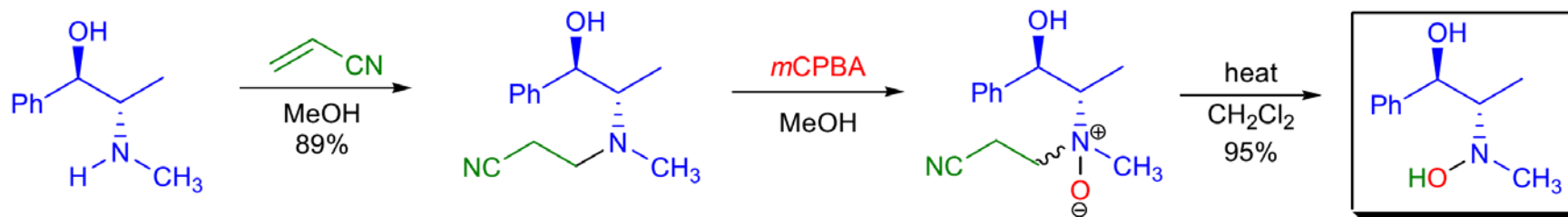
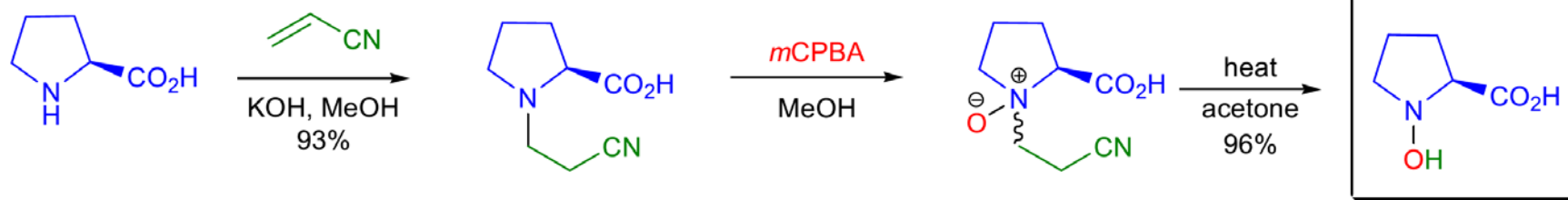
Generally the mechanism of the Hofmann elimination is E2, and it is an antielimination (the leaving groups have to be trans-diaxial/antiperiplanar).



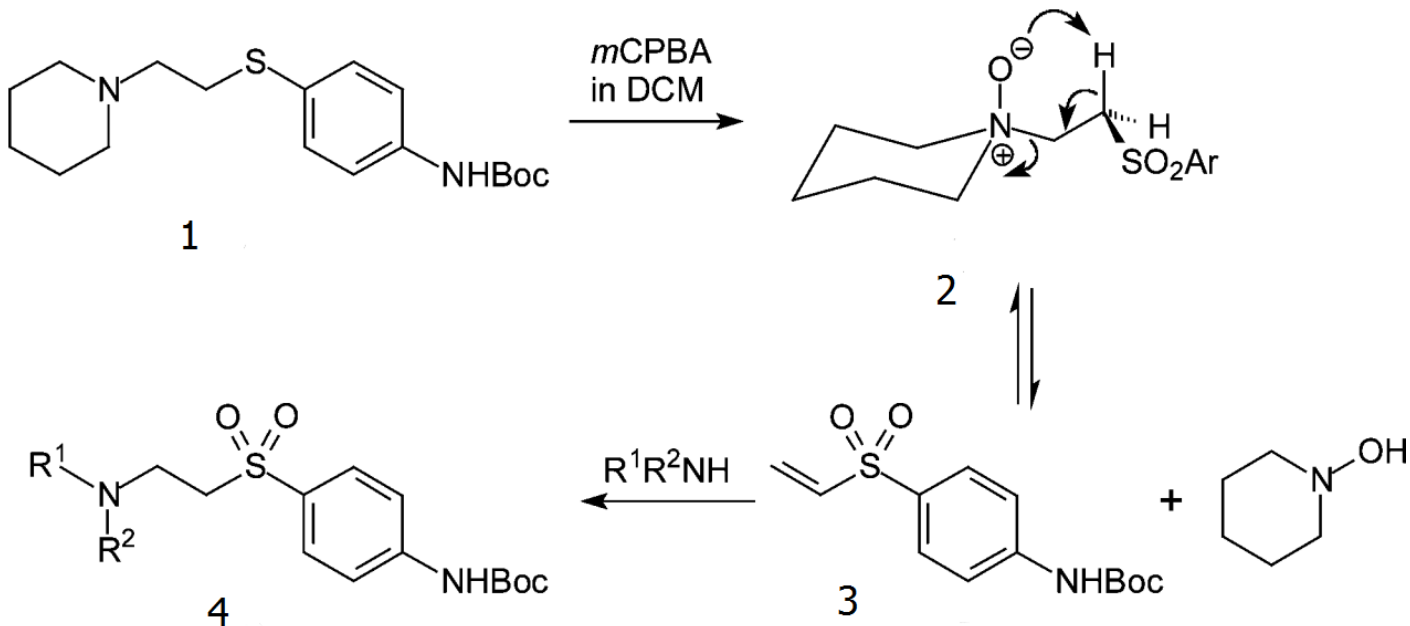
Synthetic Applications:



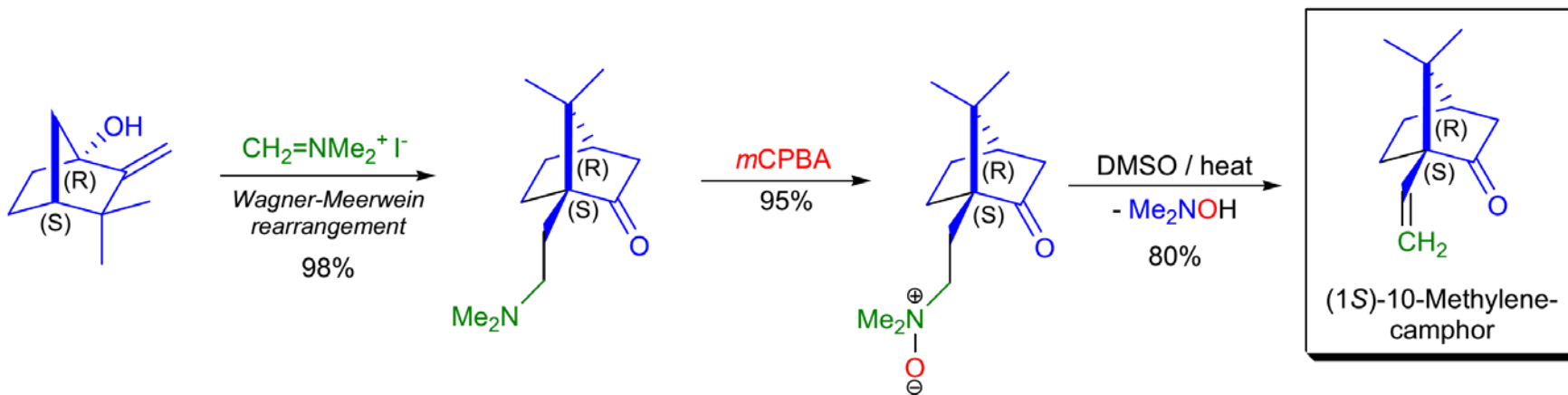
Helv. Chim. Acta **2002**, *85*, 1118-1127



Tetrahedron Lett. **2001**, *42*, 8247-8249



J. Am. Chem. Soc. **2006**, *128*, 6012-6013



Tetrahedron: Asymmetry **2002**, *13*, 17-19