Massachusetts Institute of Technology Organic Chemistry 5.512

April 6, 2005 Prof. Rick L. Danheiser

Unit 3 Stereocontrolled Conjugate Addition

- ★ Catalytic Asymmetric Conjugate Addition I: Unstabilized Nucleophiles
 Hayashi-Miyaura Rh-Catalyzed Conjugate Addition Reactions
 Cu-Catalyzed Conjugate Addition of Organozinc Compounds
 Organocatalytic Conjugate Addition of Activated Aromatic Compounds
- ★ Catalytic Asymmetric Conjugate Addition II: Conjugate Reduction
- ★ Catalytic Asymmetric Conjugate Addition III: Stabilized Nucleophiles Michael Reactions Using Heterobimetallic Catalysts (Shibasaki) Organocatalytic Michael Reactions (Jorgensen)

Organocatalytic Conjugate Addition of Activated Aromatic Compounds

Reviews: See Berkessel and Gröger

R CHO
$$\begin{array}{c}
R_2*NH_2*CI^{-} \\
-H_2O
\end{array}$$

$$\begin{array}{c}
Ph \\
N \\
H
\end{array}$$

$$\begin{array}{c}
R_2*NH_2*CI^{-} \\
NR_2*
\end{array}$$

$$\begin{array}{c}
R_2*NH_2*CHO
\end{array}$$

$$\begin{array}{c}
R_2*NH
\end{array}$$

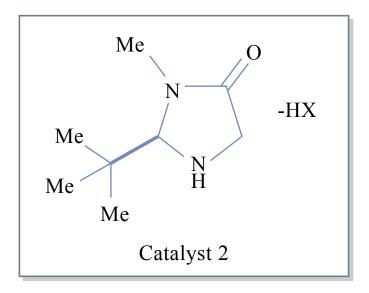
$$\begin{array}{c}
R_2*NH
\end{array}$$

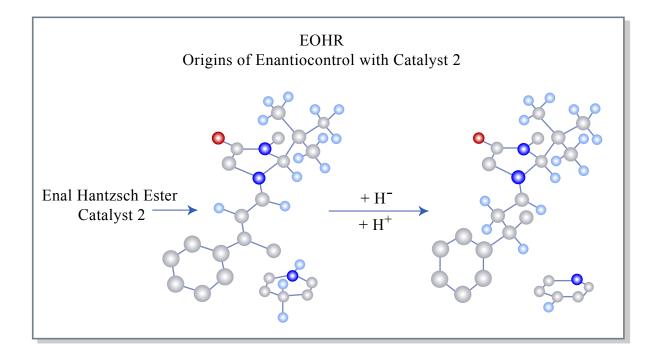
Catalytic Asymmetric Conjugate Reduction

Buchwald Cu-Catalyzed Conjugate Reduction: J. Am. Chem. Soc. 2003, 125, 11,253 and refs cited

MacMillan Organocatalytic Conjugate Reduction: J. Am. Chem. Soc. 2005, 127, 32

Figures by MIT OCW.





Figures by MIT OCW.

Catalytic Asymmetric Conjugate Addition of Stabilized Nucleophiles

I. Shibasaki: "Heterobimetallic Catalysts

II. Jorgensen: Organocatalytic Conjugate Addition Angew. Chem. Int. Ed. 2003, 42, 661 and 4955

- III. Jacobsen: Enantioselective Conjugate Additions Catalyzed by (Salen)Al Complexes *J. Am. Chem. Soc.* **2005**, *127*, 1313
- IV. Corey: Organocatalytic Conjugate Addition of Imino Esters *Org. Lett.* **2000**, *2*, 1097.

$$\begin{array}{c} \text{C}_{\text{e}}\text{H}_{5} \\ \text{O} \\ \text{O} \\ \text{F} \\ \text{O} \\ \text{O} \\ \text{F} \\ \text{O} \\ \text{C} \\ \text$$

1a $R_1 = CH_2CH = CH_2$, $R_2 = H_2C = CH$