## Massachusetts Institute of Technology Organic Chemistry 5.512

April 1, 2005 Prof. Rick L. Danheiser

# Unit 3 Stereocontrolled Conjugate Addition

- ★ Intrinisic Stereochemistry in the Michael Reaction
- ★ Substrate Control: Asymmetric Induction by Molecular Framework
- ★ Substrate Control: Asymmetric Induction via Chiral Auxiliaries
- ★ Catalytic Asymmetric Conjugate Addition I: Unstabilized Nucleophiles
- ★ Catalytic Asymmetric Conjugate Addition II: Conjugate Reduction
- ★ Catalytic Asymmetric Conjugate Addition III:

Stabilized Nucleophiles (Michael Additions)

#### **Definitions**

#### **Conjugate Addition**

Addition of a nucleophile to an alkene or alkyne conjugated with an electron-withdrawing group

#### **Michael Reaction**

Addition of an enolate or related "stabilized" carbanionic species to an  $\alpha,\beta$ -unsaturated carbonyl compound or related electron-deficient alkene or alkyne

#### **General References**

Conjugate Addition Reactions in Organic Synthesis", P. Perlmutter, Pergamon Press, 1992

#### Reviews:

"Enantioselective Conjugate Additions"; M. P. Sibi; S. Manyem *Tetrahedron* **2000**, *56*, 8033. "Recent Advances in Catalytic Enantioselective Michael Additions";

N. Krause; A. Hoffmann-Röder Synthesis 2001, 171

### **Review Reading Assignment**

Carey and Sundberg "Advanced Organic Chemistry" Part B

Section 1.10 (Alkylation of Carbon Nucleophiles by Conjugate Addition)

Section 7.3.1 (Organozinc Compounds)

Section 8.1 (Organocopper Intermediates)

Section 9.1.1 (Synthesis of Organoboranes)