## **Massachusetts Institute of Technology**

**Organic Chemistry 5.512** 

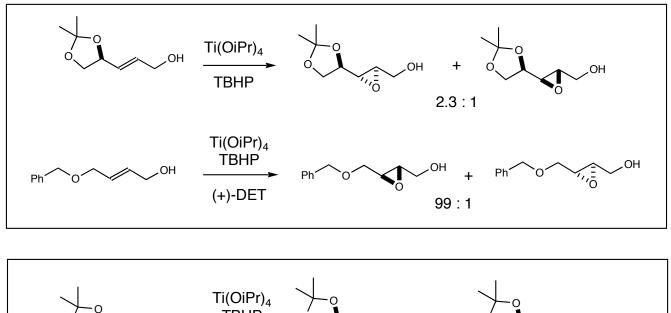
March 7, 2005 Prof. Rick L. Danheiser

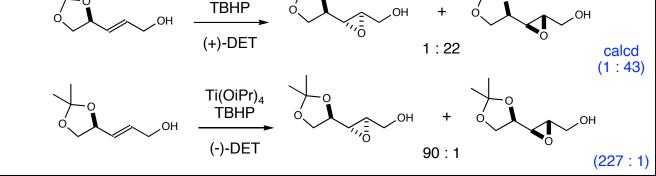
## **Strategies for Stereocontrolled Synthesis**

## st Reagent Control Strategies and Double Asymmetric Synthesis $\,st$

"What changes may organic synthesis undergo? . . . . With appropriate chiral reagents and catalysts at hand, the synthetic design of many natural (and unnatural) products will become straightforward, and as a result some of the aesthetic elements of traditional organic synthesis, as exemplified by the synthesis of erythronolide A in Section 7, may well be lost. . . . . However, the power of the new strategy has already made possible what appeared to be almost impossible even a few years ago. In this sense a new era which is characterized by the evolution from substrate-controlled to reagent-controlled organic synthesis is definitely emerging."

S. Masamune et al., "Double Asymmetric Synthesis and a New Strategy for Stereochemical Control in Organic Synthesis", *Angew. Chem. Int. Ed.* **1985**, *24*, 1.





## Comparison of Substrate and Reagent Control Strategies

	Advantages	Disadvantages
Substrate Control	$\star$ Exploits resident chirality	$\star$ Requires strong bias in substrate
		★ Different strategy needed for each epimer
Reagent Control	★ Same strategy sometimes applicable to synthesis of both epimers	★ Not applicable if substrate has strong bias
	$\star$ Applicable to substrates with low bias	★ Requires reagents with very strong bias