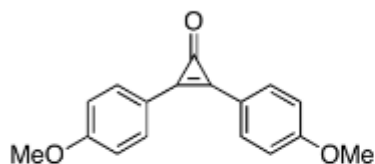
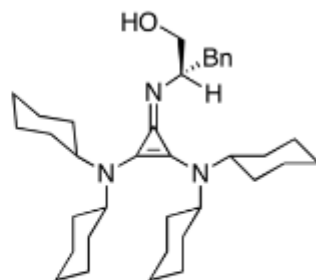




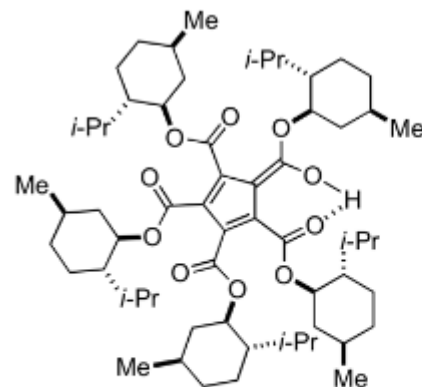
## The Career of Tristan H. Lambert



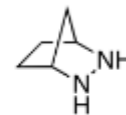
dehydrative  
catalysis



enantioselective  
Brønsted base  
catalysis



enantioselective  
Brønsted acid  
catalysis



catalytic  
carbonyl-olefin  
metathesis

Jian Rong (荣健)

Hu Group Meeting

Apr 11, 2016

# Tristan H. Lambert: Biographical Notes



born in Madison, Wisconsin in 1976

## Education

**2004-2006:** NIH Postdoctoral Fellow, Memorial Sloan-Kettering Cancer Center (Samuel J. Danishefsky, advisor)

**2004:** Ph.D. Chemistry, California Institute of Technology (David W. C. MacMillan, advisor)

**2000:** M.S. Chemistry, University of California at Berkeley (David W. C. MacMillan, advisor)

**1998:** B.S. Chemistry, University of Wisconsin at Platteville

## Professional experience

**2011-present:** Associate Professor, Columbia University

**2006-2011:** Assistant Professor, Columbia University

**2004-2006:** NIH Postdoctoral Fellow, Memorial Sloan-Kettering Cancer Center

**1998-2003:** Graduate Research Assistant, UC-Berkeley and Caltech

**1997-1998:** Undergraduate Research, UW-Platteville

**Summer 1997:** NSF--Research Experience for Undergraduates, University of Kansas

## Awards And Honors

**2004-2006** NIH Postdoctoral Fellowship

**2009** Abbott Young Investigator Award

**2010** NSF CAREER award

**2010** Thieme SynLett/Synthesis Journal Award

**2010** Alfred P. Sloan Research Fellow

**2010** Amgen Young Investigator Award

**2010** ACS Young Investigators Symposium

**2011** Eli Lilly Grantee Award

**2014** Visiting Professorship, Phillips-University Marburg

**2015** Swiss Chemical Society Lecturer

# Lambert Group: Members and Publications

## Current Members



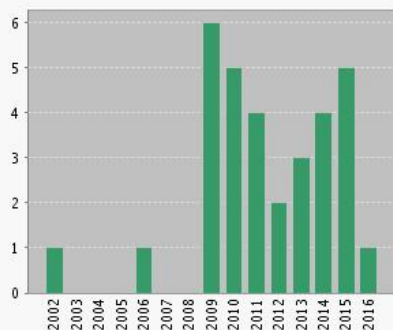
### 引文报告: 32

(来自 所有数据库)

您的检索: 作者: (Lambert Tristan) ...[更多内容](#)

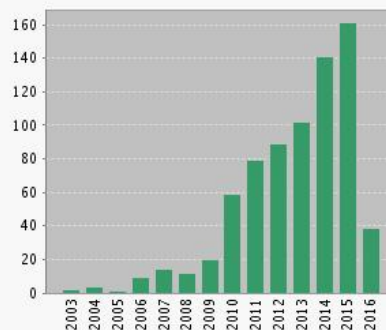
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每年出版的文献数

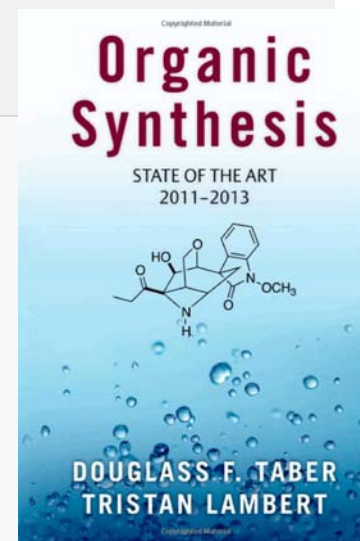


显示最近 20 年。

每年的引文数



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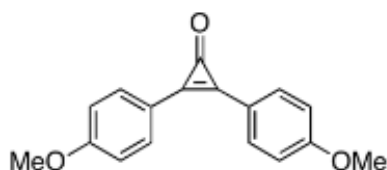


# Significant Research Areas

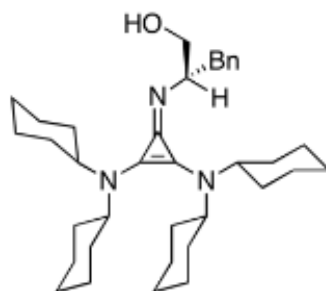


*Reaction Design | Synthesis*

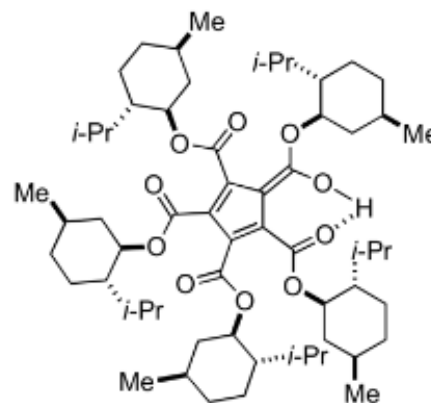
novel catalytic strategies for  
selective organic synthesis



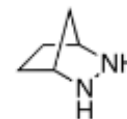
dehydrative  
catalysis



enantioselective  
Brønsted base  
catalysis

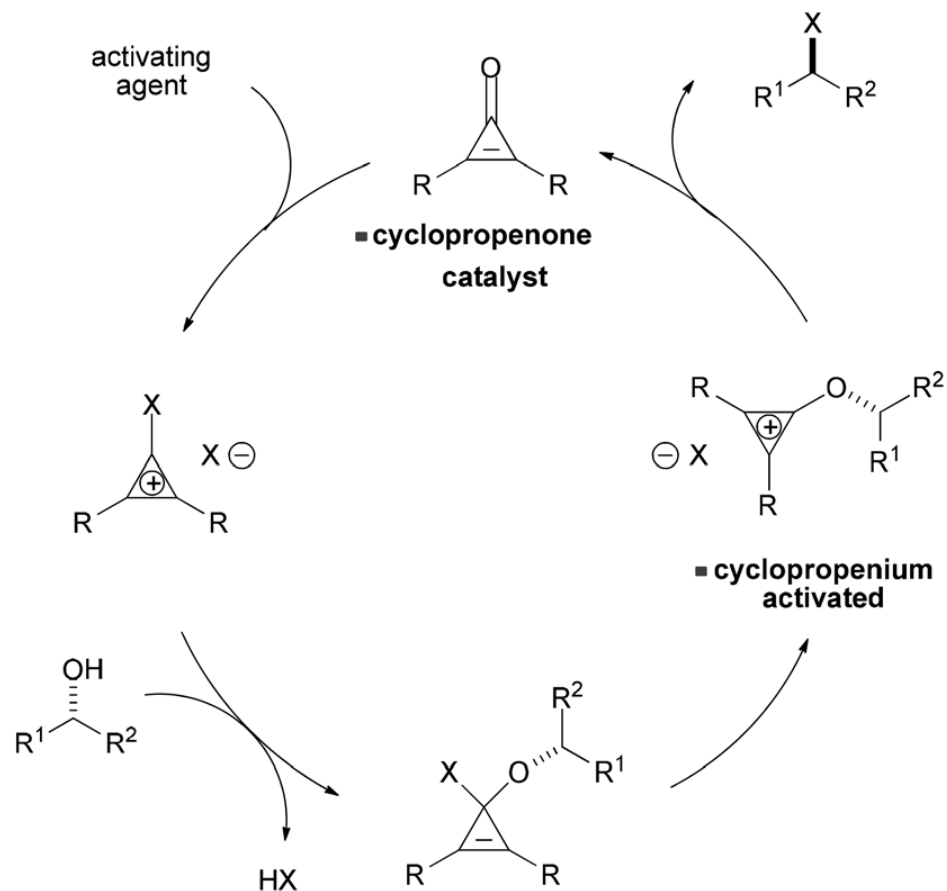


enantioselective  
Brønsted acid  
catalysis



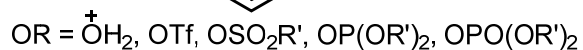
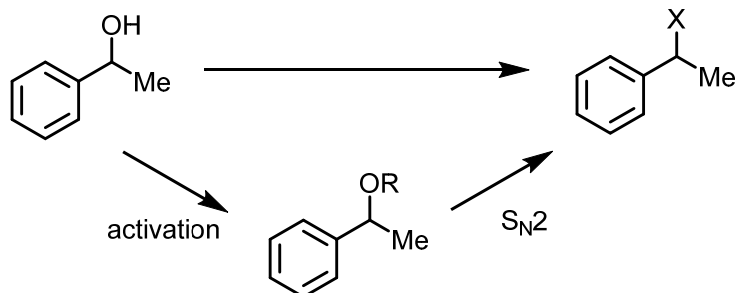
catalytic  
carbonyl-olefin  
metathesis

# Dehydrative catalysis (Aromatic cation activation)

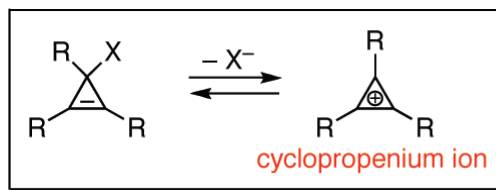


# Dehydrative catalysis (Aromatic cation activation)

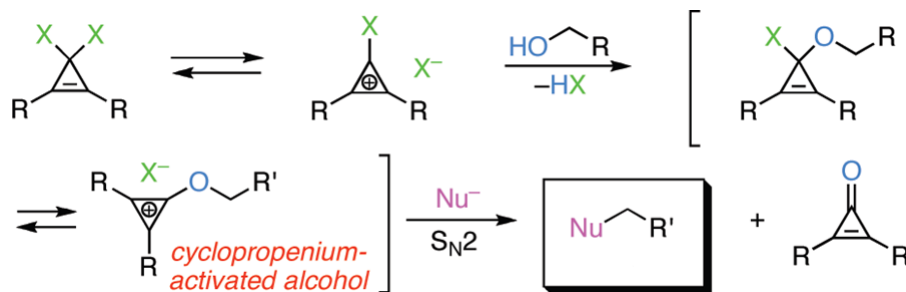
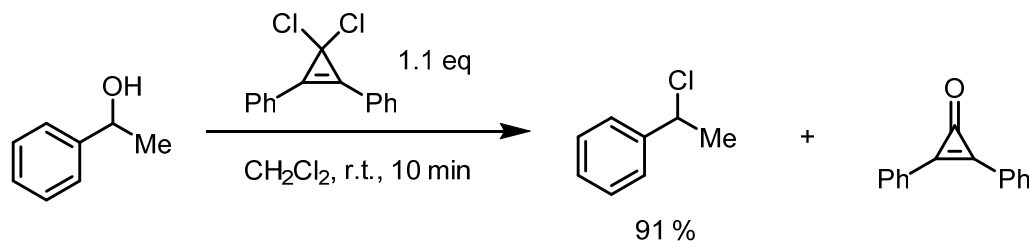
## Aromatic Cation Activation of Alcohols



### Activation of Alcohols

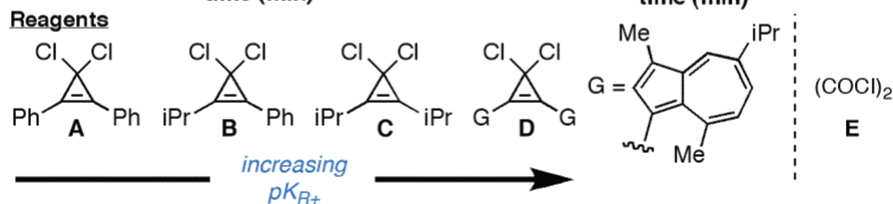
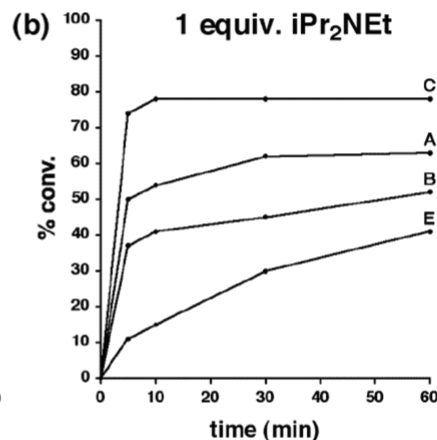
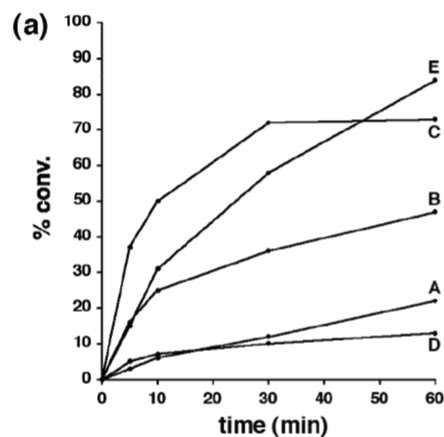
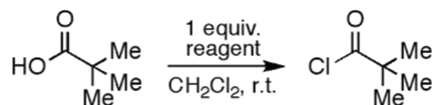
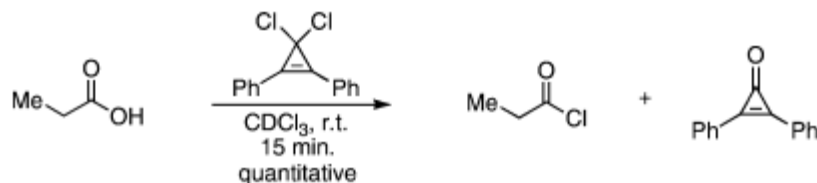


- $2\pi$ -electron aromatic system
- discovered by Breslow (1957)
- highly stabilized carbocations
- electronically, sterically tunable
- potential for new reaction design



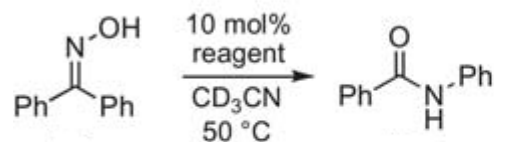
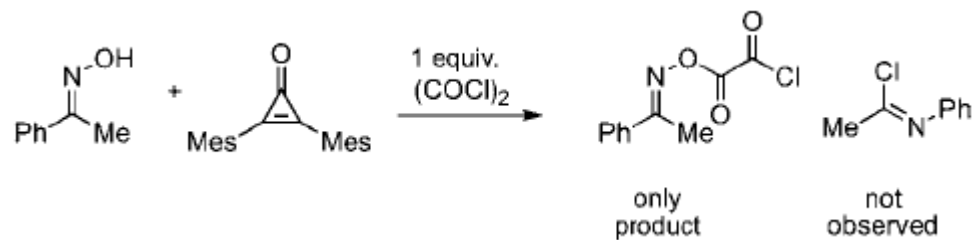
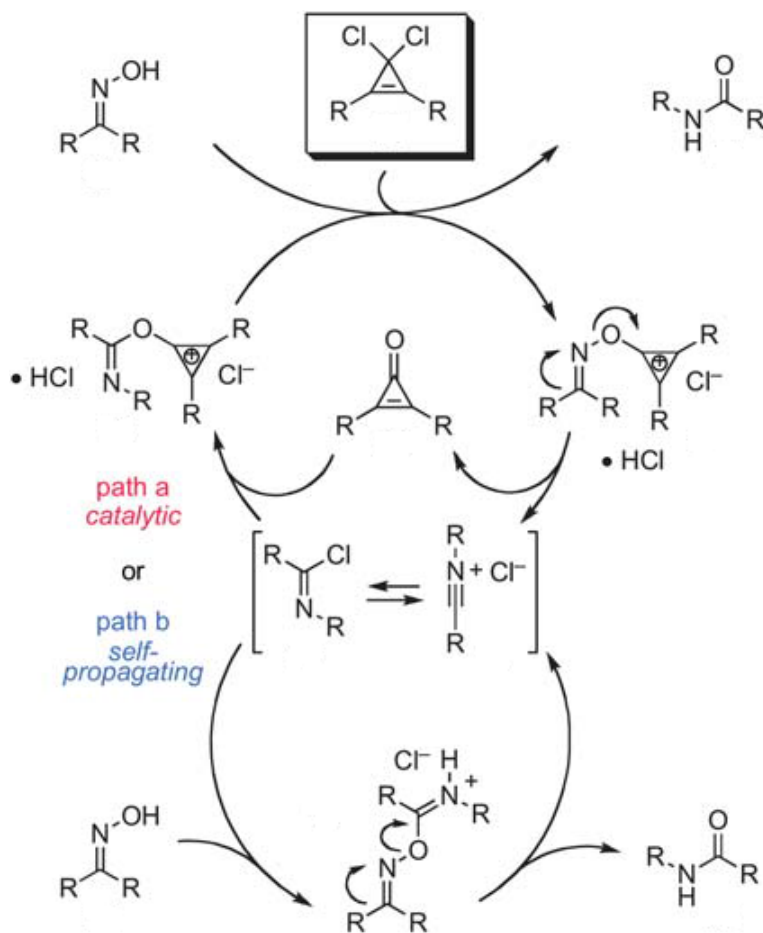
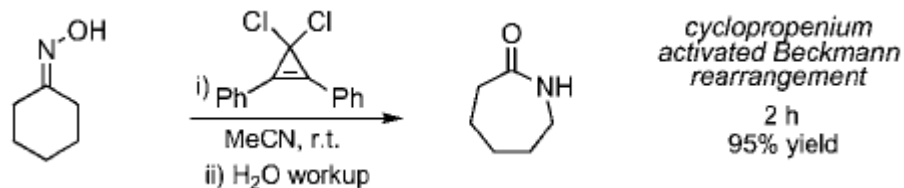
# Dehydrative catalysis (Aromatic cation activation)

## Aromatic Cation Activation of Carboxylic Acids: Rapid Generation of Acid Chlorides

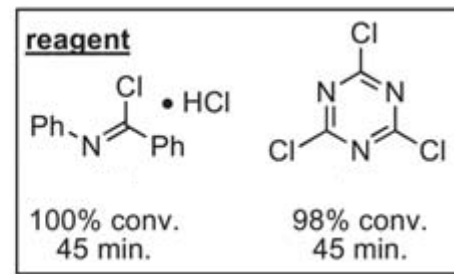


# Dehydrative catalysis (Aromatic cation activation)

## Cyclopropenium-activated Beckmann rearrangement

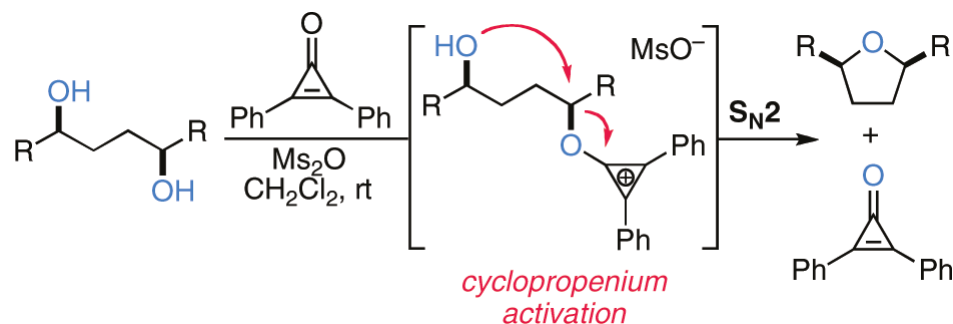
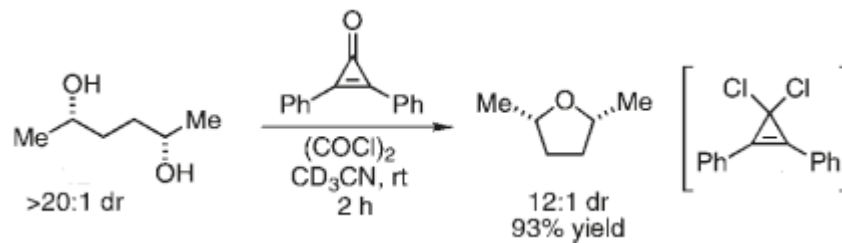


same rate



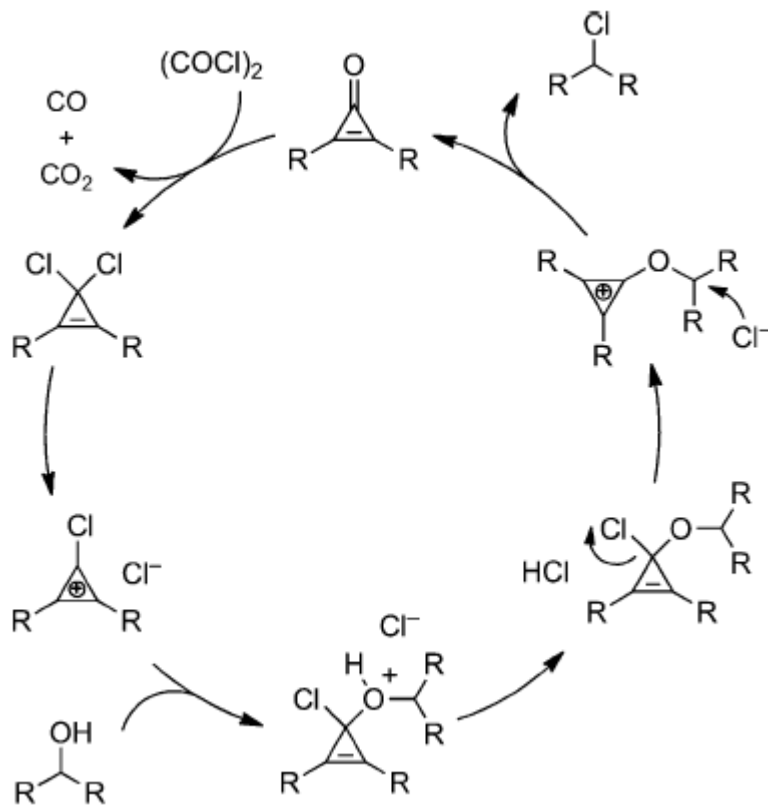
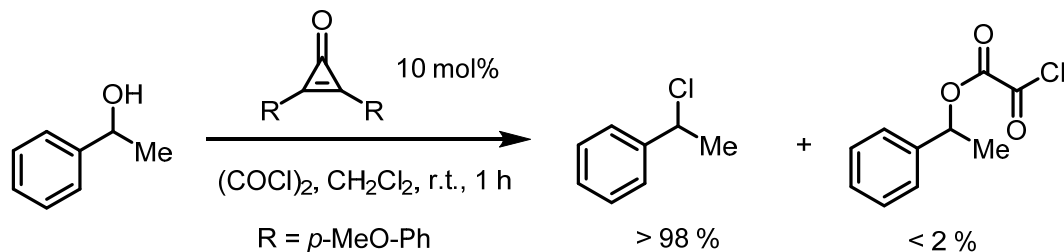
# Dehydrative catalysis (Aromatic cation activation)

## Cyclopropenium-Activated Cyclodehydration of Diols



# Dehydrative catalysis (Aromatic cation activation)

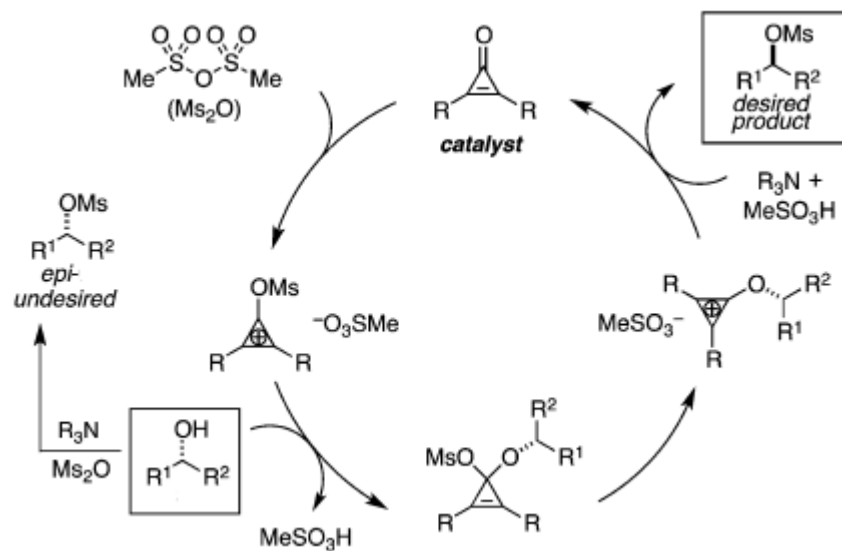
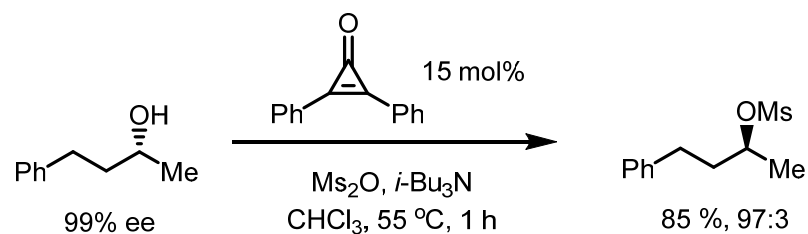
## Cyclopropenone-Catalyzed Chlorodehydration of Alcohols



The more electron-rich catalysts improved this ratio.

# Dehydrative catalysis (Aromatic cation activation)

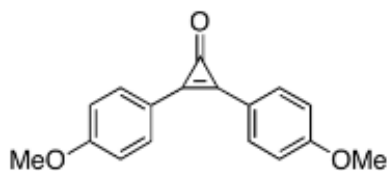
## Cyclopropenone-Catalyzed Substitution of Alcohols with Mesylate Ion



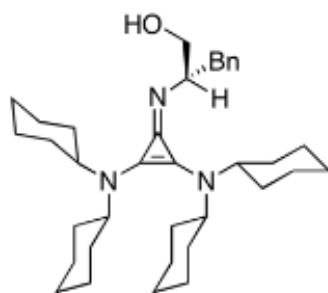
# Significant Research Areas



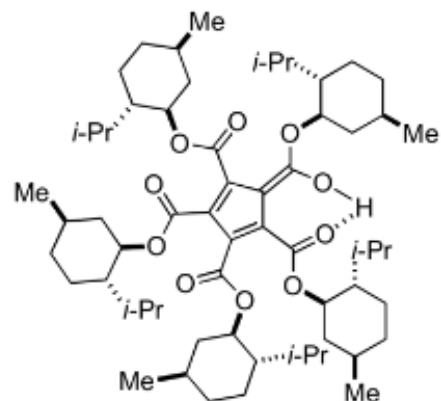
Reaction Design | Synthesis



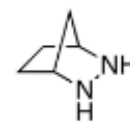
dehydrative  
catalysis



enantioselective  
Brønsted base  
catalysis



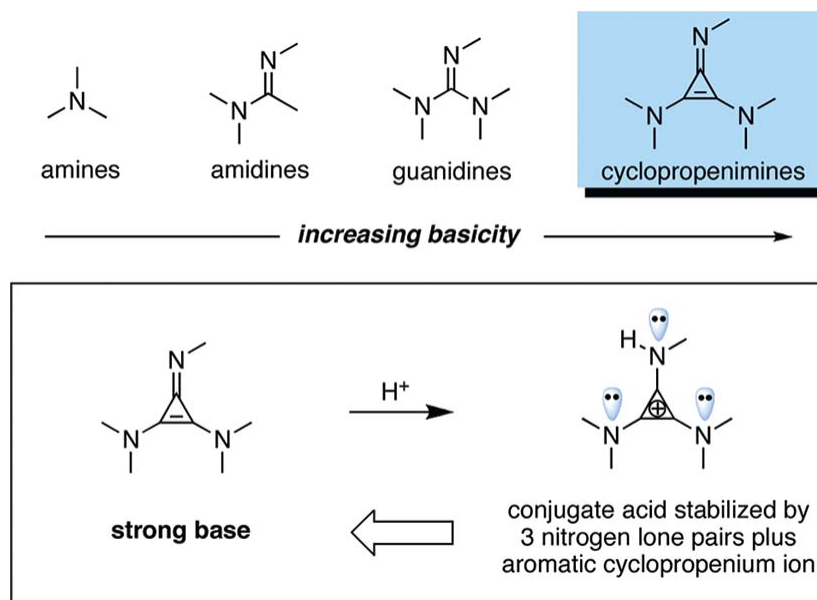
enantioselective  
Brønsted acid  
catalysis



catalytic  
carbonyl-olefin  
metathesis

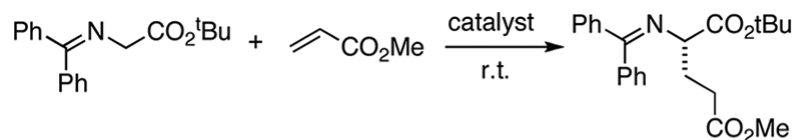
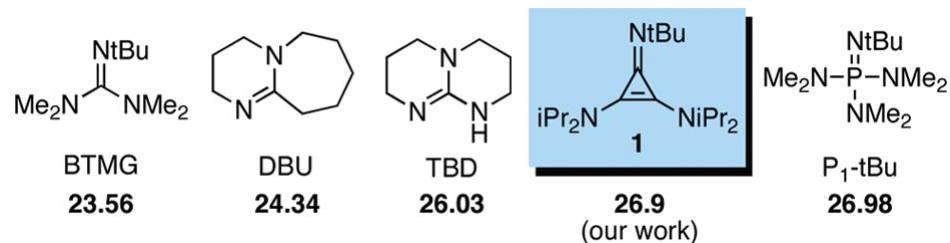
# Enantioselective Brønsted base catalysis

## Enantioselective Brønsted Base Catalysis with Chiral Cyclopropenimines

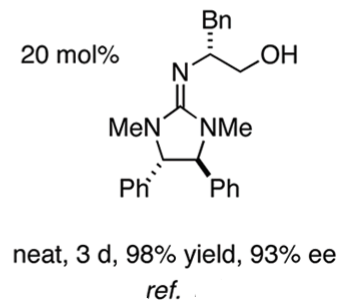
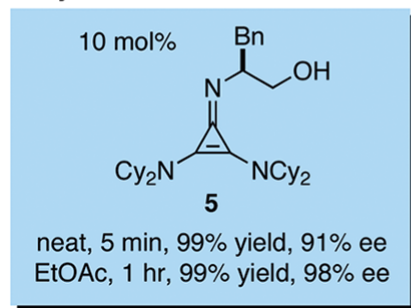


# Enantioselective Brønsted base catalysis

## Enantioselective Brønsted Base Catalysis with Chiral Cyclopropenimines

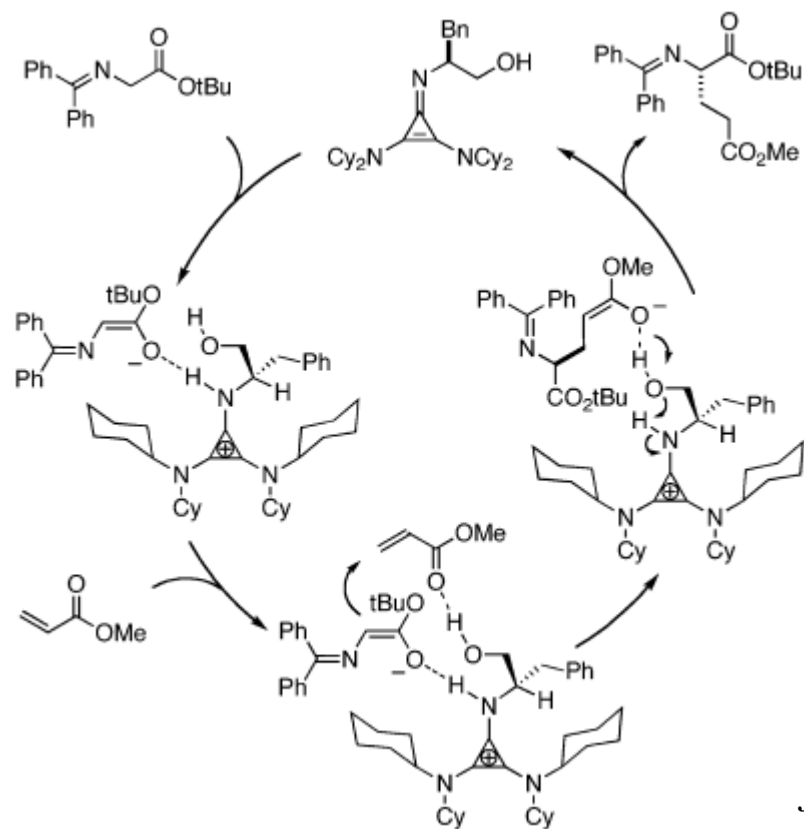
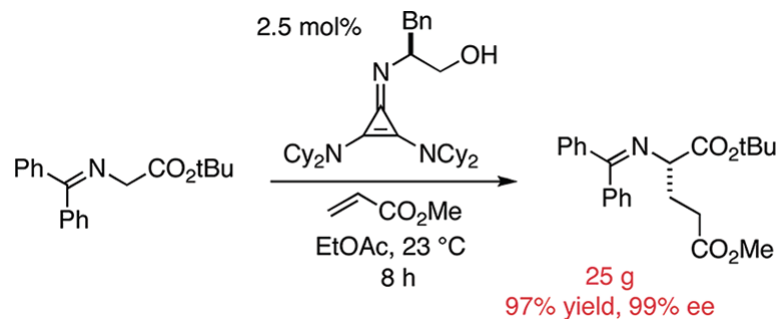


Catalysts:



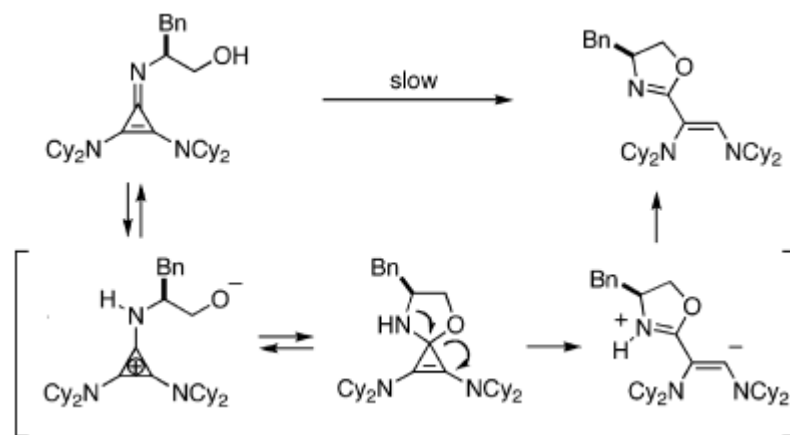
# Enantioselective Brønsted base catalysis

## Enantioselective Brønsted Base Catalysis with Chiral Cyclopropenimines



# Enantioselective Brønsted base catalysis

## Enantioselective Brønsted Base Catalysis with Chiral Cyclopropenimines



cyclopropenimine

$t_{1/2} \approx 12$  days at r.t.

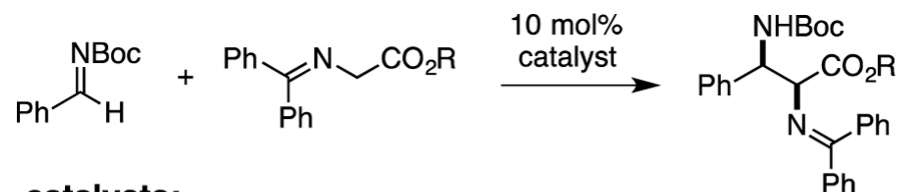
Storing should be at  $-20$  °C, and 94% remained after 30 days.

HCl salt is indefinitely stable at rt.

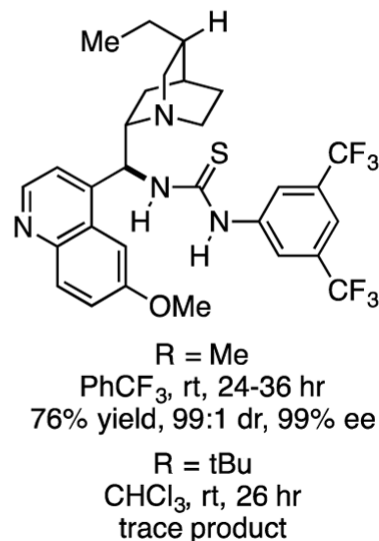
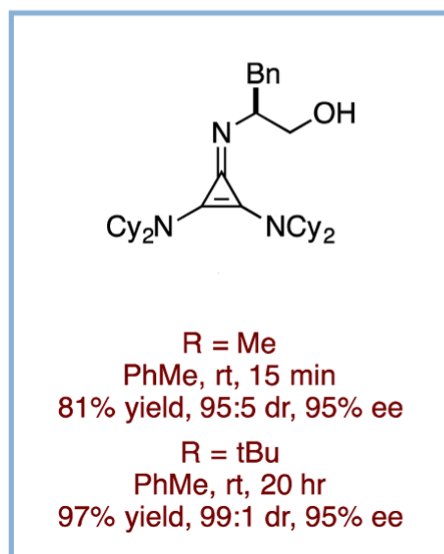
It's convenient to store the catalyst as its acid co-salt.

# Enantioselective Brønsted base catalysis

## Cyclopropenimine-Catalyzed Enantioselective Mannich Reactions of *tert*-Butyl Glycinates with *N*-Boc-Imines

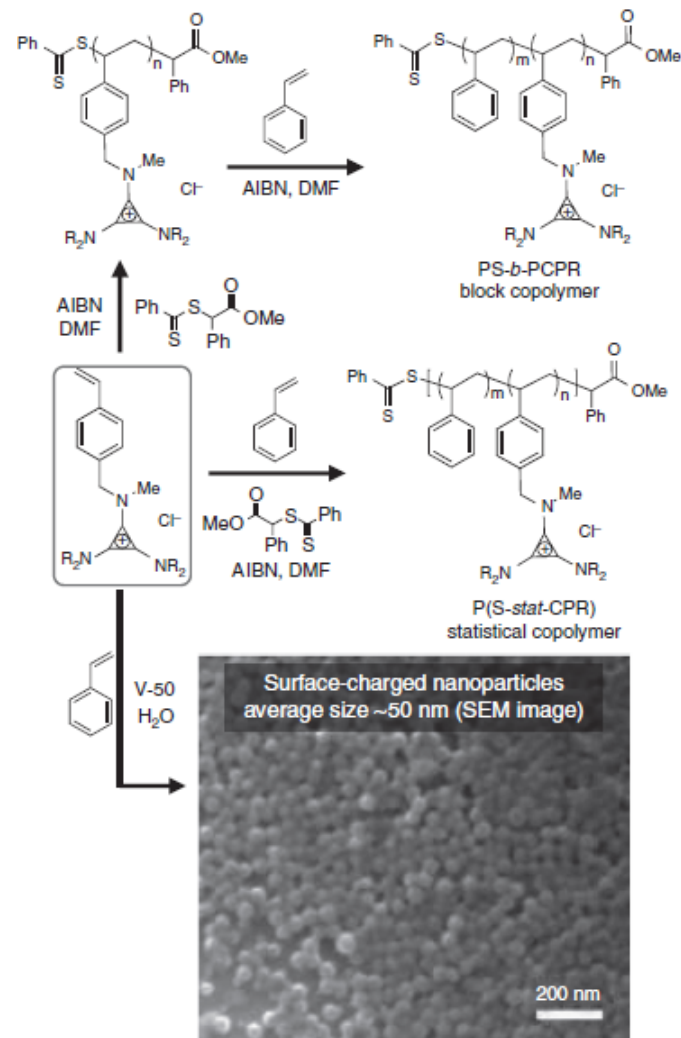
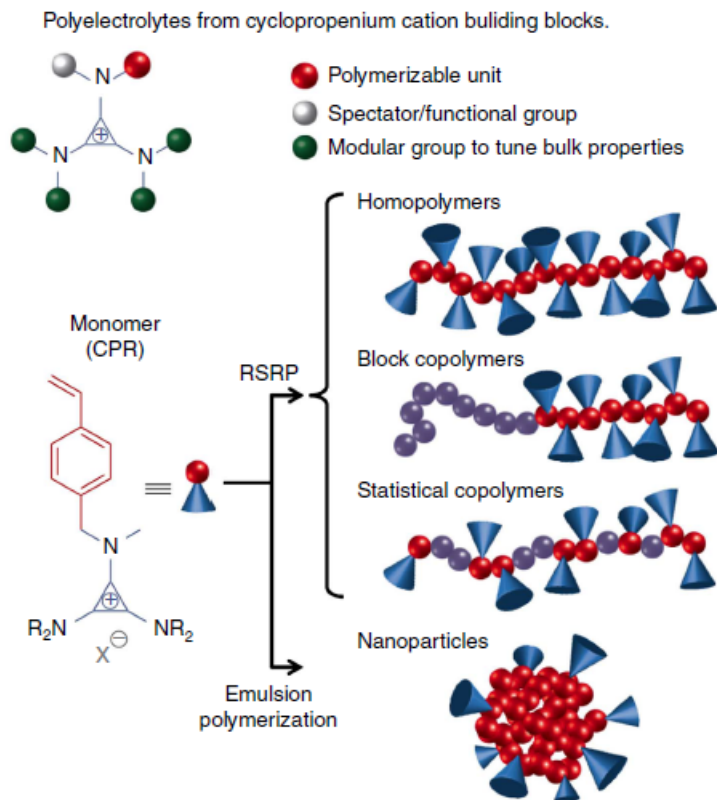


**catalysts:**



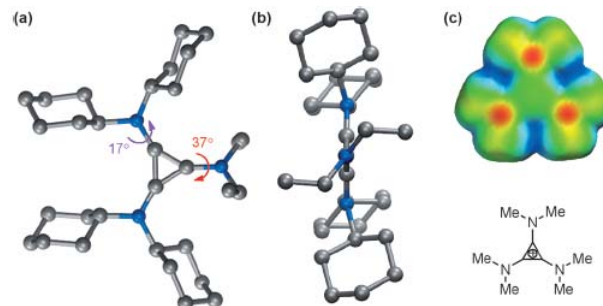
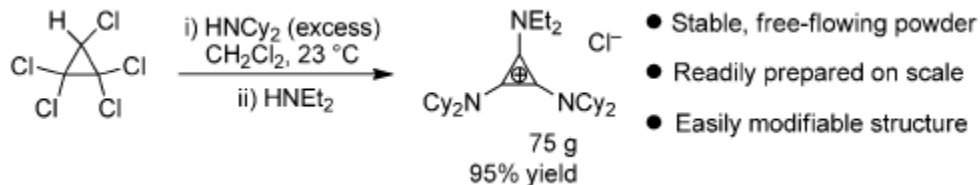
# Enantioselective Brønsted base catalysis

## The evolution of cyclopropenium ions into functional polyelectrolytes

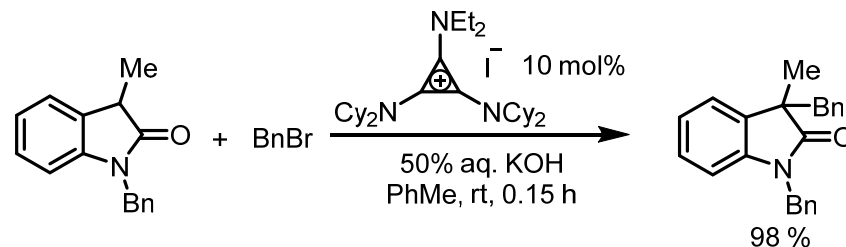
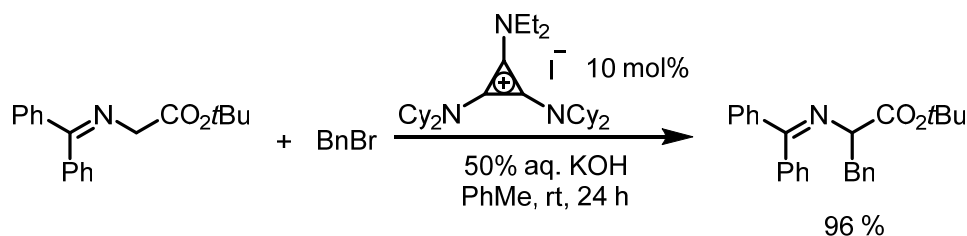


# Enantioselective Brønsted base catalysis

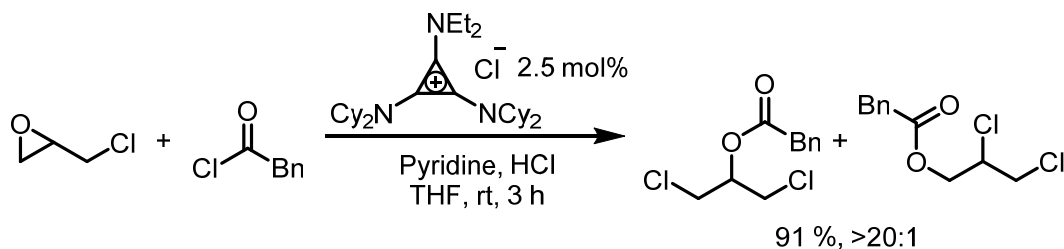
## Phase-Transfer and Other Types of Catalysis with Cyclopropenium Ions



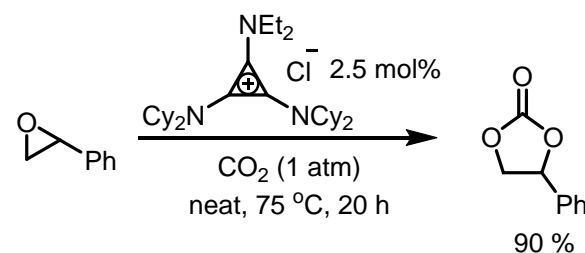
### PTC of enolate alkylation



### Addition of acid chlorides to epoxides

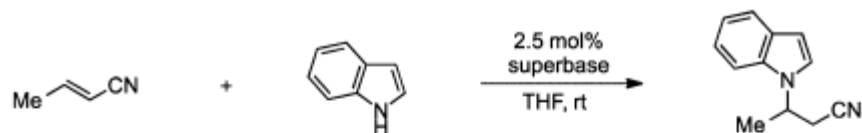


### Addition of carbon dioxide to epoxides

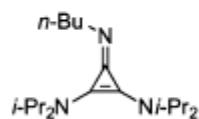


# Enantioselective Brønsted base catalysis

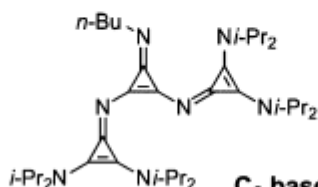
## Higher-Order Cyclopropenimine Superbases: Direct Neutral Brønsted Base Catalyzed Michael Reactions with $\alpha$ -Aryl Esters



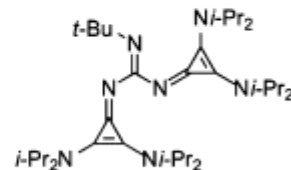
superbase:



**C<sub>1</sub> base**  
pK<sub>BH<sup>+</sup></sub> = 27.6  
0% conv.  
24 h



**C<sub>3</sub> base**  
pK<sub>BH<sup>+</sup></sub> = 31.6  
89% yield  
3 h

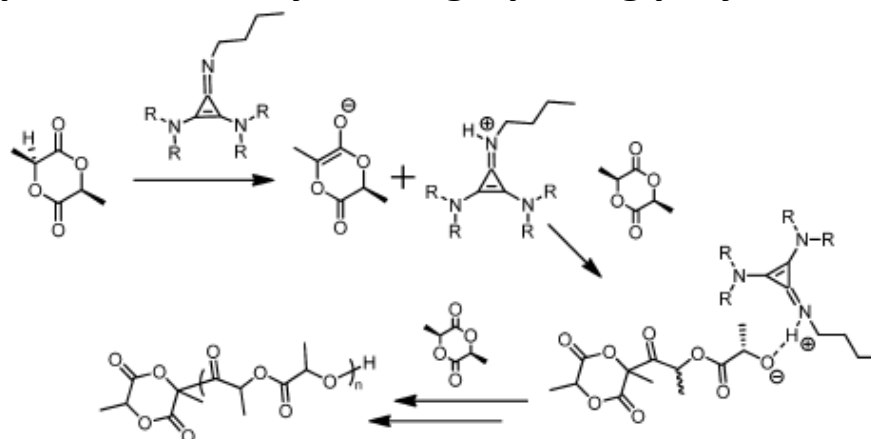


**GC<sub>2</sub> base**  
pK<sub>BH<sup>+</sup></sub> = 35.6  
95% yield  
3 h

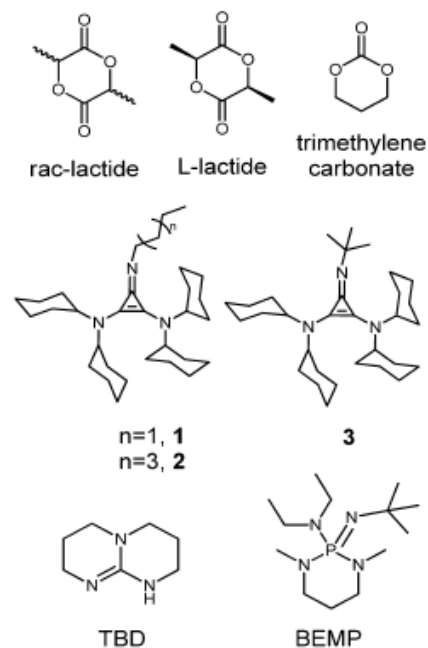
enable reactivity with a broader range of less acidic substrates

# Enantioselective Brønsted base catalysis

## Cyclopropenimine Superbases-catalyzed ring-opening polymerization



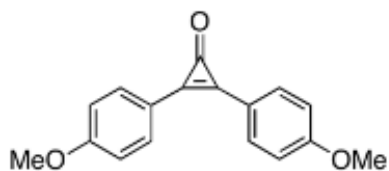
entry	monomer	catalyst	cat. conc. (M)	solvent	time	conv. (%)	$M_n^b$	$M_w/M_n$
1	rac-lactide	1	0.010	CH <sub>2</sub> Cl <sub>2</sub>	8 min	84	8390	1.42
2	rac-lactide	1	0.050	CH <sub>2</sub> Cl <sub>2</sub>	10 min	99	15300	1.57
3 <sup>c</sup>	rac-lactide	1	0.007	C <sub>6</sub> D <sub>6</sub>	2 days	99	70700	1.46
4	L-lactide	1	0.010	CH <sub>2</sub> Cl <sub>2</sub>	8 min	93	11500	1.30
5	L-lactide	2	0.010	CH <sub>2</sub> Cl <sub>2</sub>	8 min	90	17300	1.46
6	rac-lactide	3	0.020	CD <sub>2</sub> Cl <sub>2</sub>	20 min	98	13100	1.38
7	rac-lactide	BEMP	0.010	CH <sub>2</sub> Cl <sub>2</sub>	8 min	65	8300	1.24
8	L-lactide <sup>d</sup>	TBD	0.0007	CH <sub>2</sub> Cl <sub>2</sub>	10 s	11	19000	1.39
9	carbonate <sup>e</sup>	1	0.010	CH <sub>2</sub> Cl <sub>2</sub>	22 hrs	3	-	-



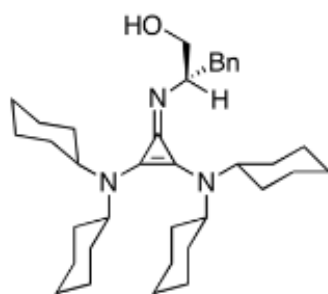
# Significant Research Areas



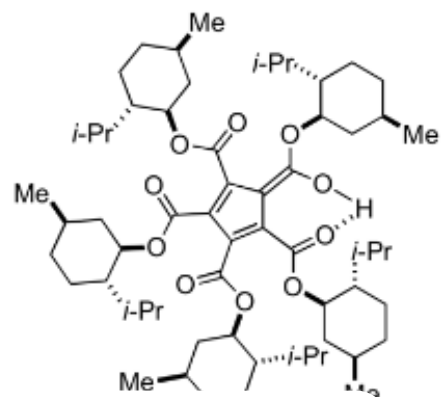
Reaction Design | Synthesis



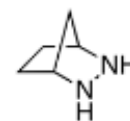
dehydrative  
catalysis



enantioselective  
Brønsted base  
catalysis



enantioselective  
Brønsted acid  
catalysis

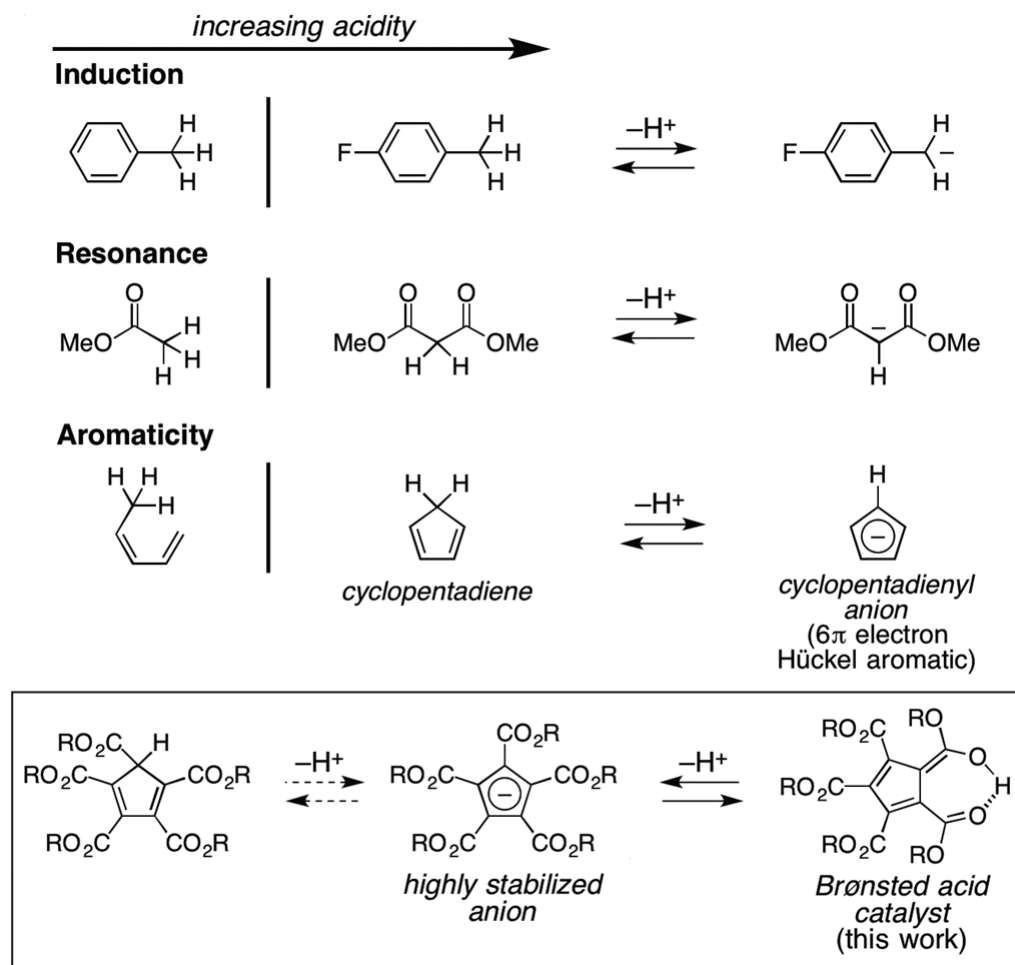


catalytic  
carbonyl-olefin  
metathesis

# Enantioselective Brønsted acid catalysis

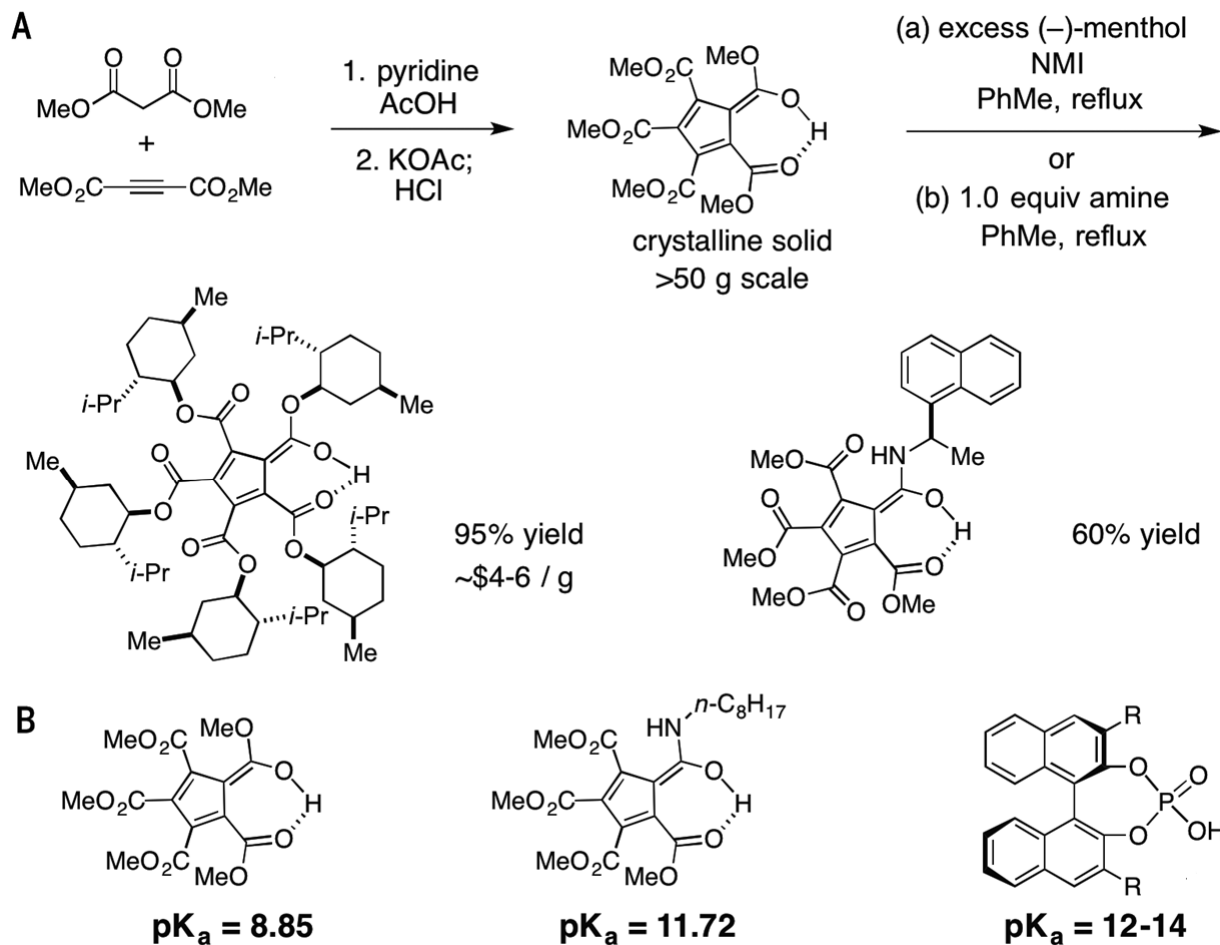
## An aromatic ion platform for enantioselective Brønsted acid catalysis

### Tuning carbon acids for catalysis



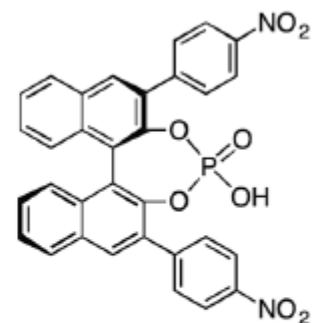
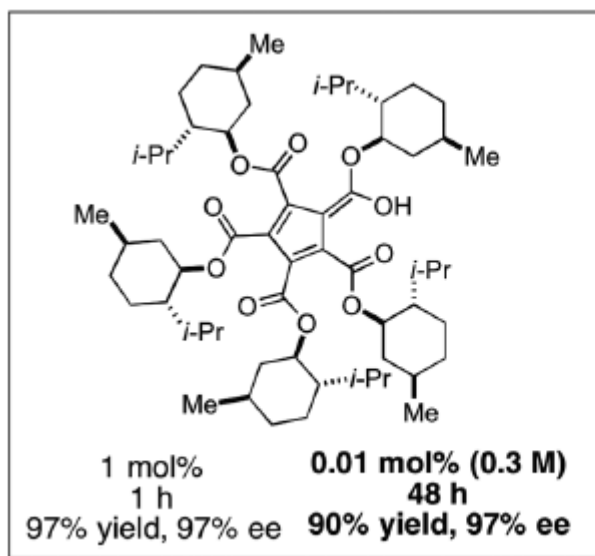
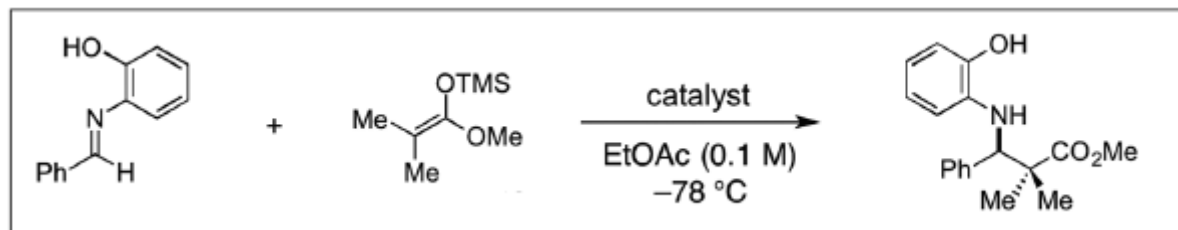
# Enantioselective Brønsted acid catalysis

## An aromatic ion platform for enantioselective Brønsted acid catalysis



# Enantioselective Brønsted acid catalysis

An aromatic ion platform for enantioselective Brønsted acid catalysis

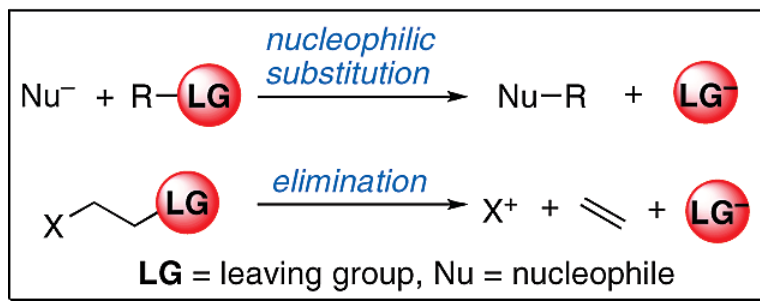


10 mol%, 24 h  
98% yield, 89% ee  
(toluene)

# Enantioselective Brønsted acid catalysis

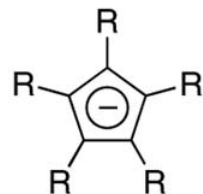
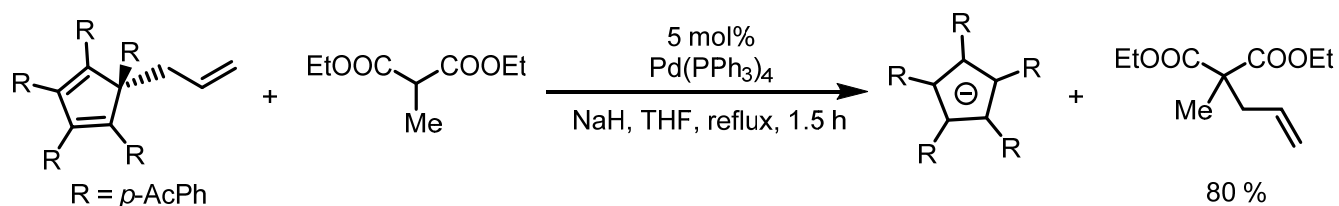
## Leaving Group Potential of a Substituted Cyclopentadienyl Anion Toward Oxidative Addition

### Leaving groups in organic chemistry



- Concept of leaving groups intrinsic to many reactions
- Typically LG =  $\text{Cl}^-$ ,  $\text{Br}^-$ ,  $\text{I}^-$ ,  $\text{RSO}_3^-$ ,  $\text{H}_2\text{O}$ ,  $\text{ROH}$ ,  $\text{R}_3\text{N}$
- $\text{R}_3\text{C}^-$  usually very poor LG

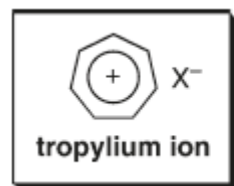
### Carbon as a leaving group



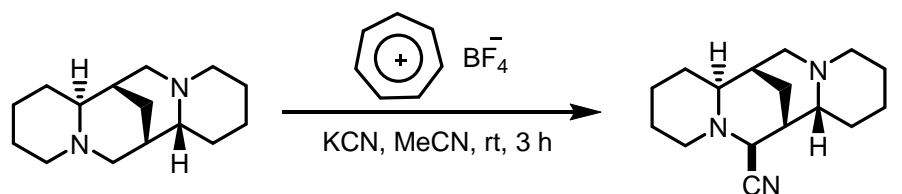
- Highly stabilized carbanions
- pKa of conjugate acids as low as -10
- Readily synthesized
- Electronically and sterically tunable
- General carbon leaving group?

# Enantioselective Brønsted acid catalysis

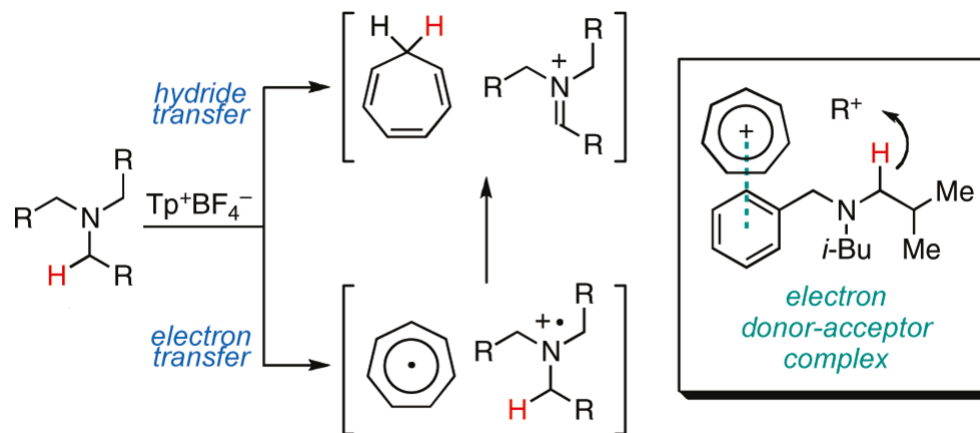
## Tropylium Ion Mediated $\alpha$ -Cyanation of Amines



- 6 $\pi$ -electron aromatic system
- first prepared by Doering and Knox in 1954
- aromatic carbocation



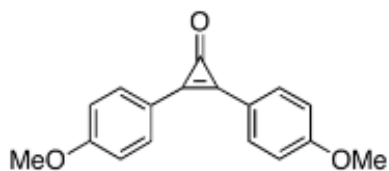
90 %, 1g scale  
R-cyanate (-)sparteine



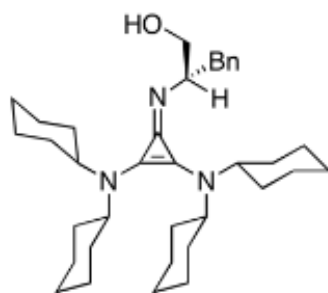
# Significant Research Areas



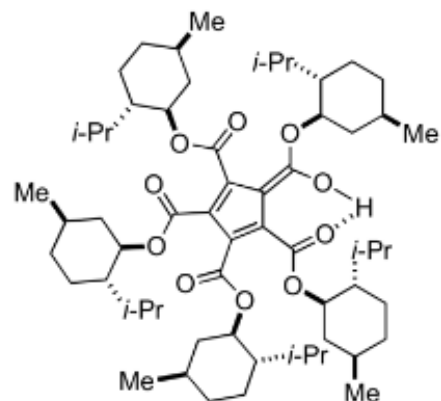
Reaction Design | Synthesis



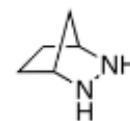
dehydrative  
catalysis



enantioselective  
Brønsted base  
catalysis



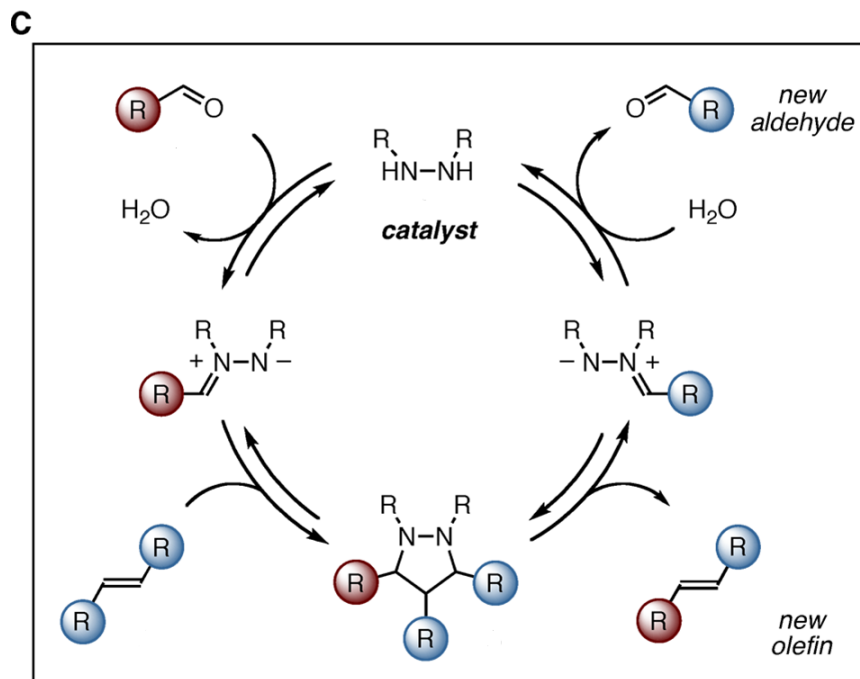
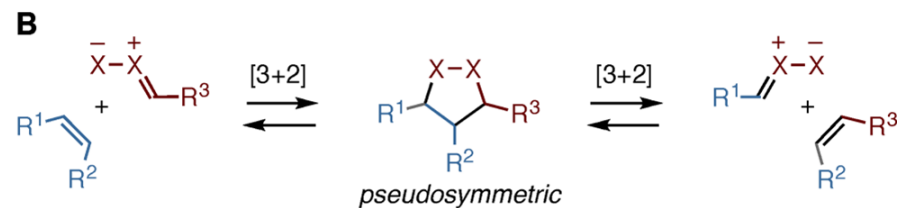
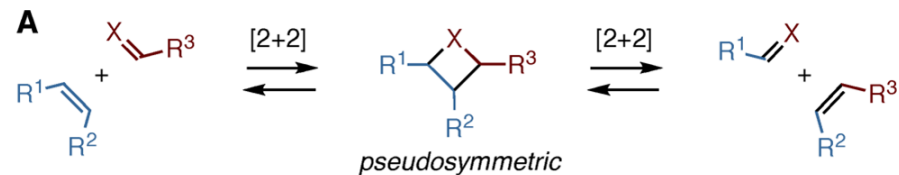
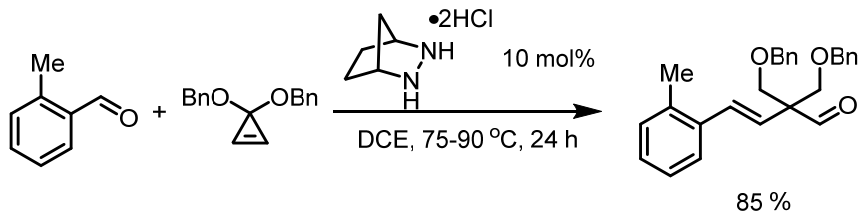
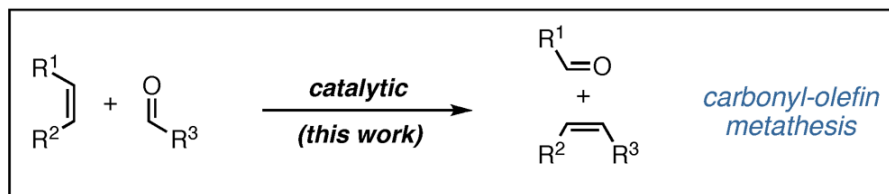
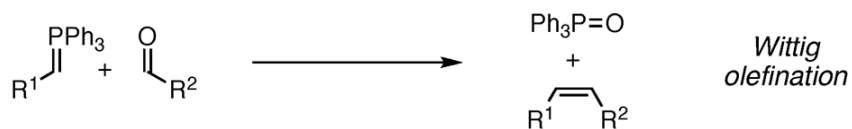
enantioselective  
Brønsted acid  
catalysis



catalytic  
carbonyl-olefin  
metathesis

# Catalytic carbonyl-olefin metathesis

## Organocatalytic Carbonyl-Olefin Metathesis

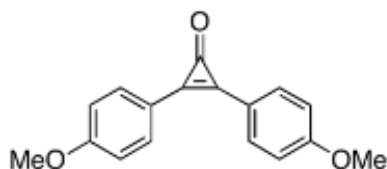


# Summary

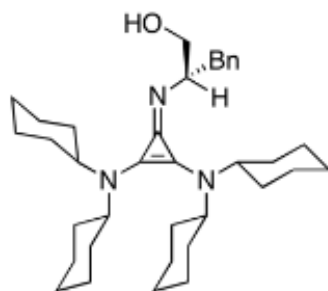


Reaction Design | Synthesis

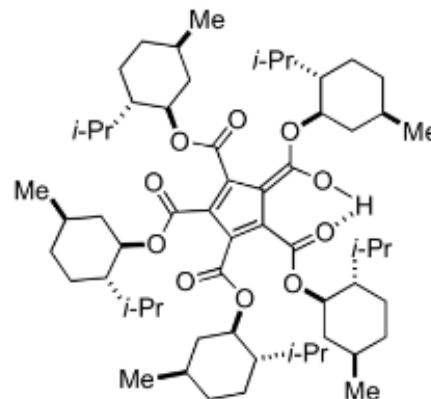
## Aromatic ions activation



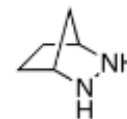
dehydrative  
catalysis



enantioselective  
Brønsted base  
catalysis



enantioselective  
Brønsted acid  
catalysis

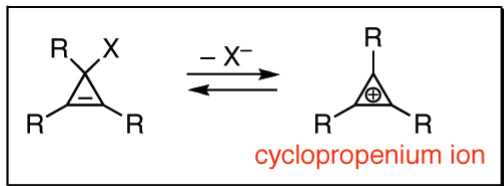


catalytic  
carbonyl-olefin  
metathesis

## Aromatic Cation

## Aromatic Cation

## Aromatic anion



- 2π-electron aromatic system
- discovered by Breslow (1957)
- highly stabilized carbocations
- electronically, sterically tunable
- potential for new reaction design

