

One- and Two –Component Domino reactions

Dong group at UT Austin

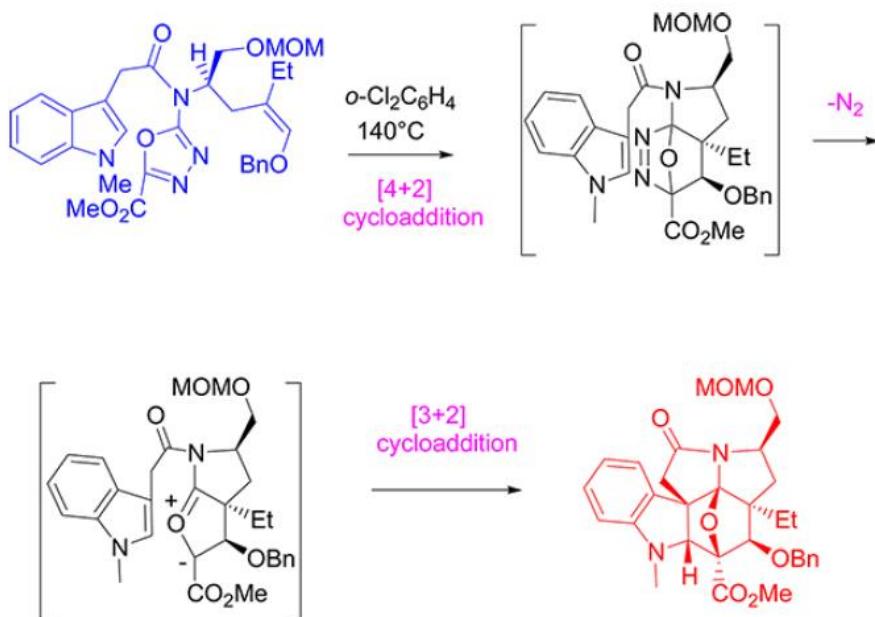
Xuan Zhou

04/15/2015

CHEMICAL REVIEWS

Stereocontrolled Domino Reactions

Hélène Pellissier*



Chem. Rev. **2013**, *113*, 442–524

One- and Two –Component Domino reactions

Domino reaction: two or more bond-forming in one reaction vessel, and no additional reagents, catalysts, or additives can be added during the reaction.

Content

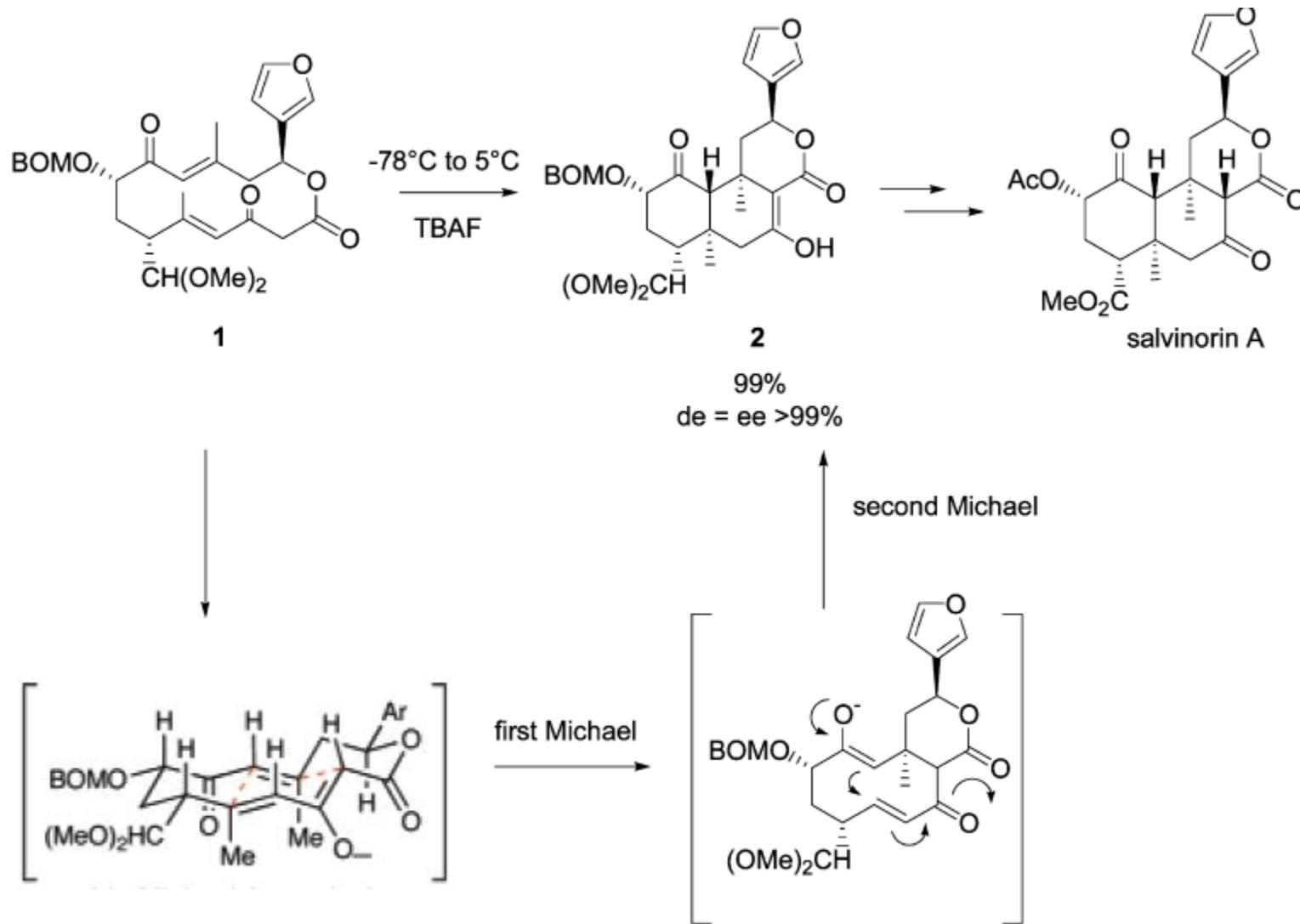
- A, Anionic primary step
- B, Cationic sequence
- C, Initiated by a pericyclic primary step
- D, Carbene sequence
- E, Palladium catalyzed domino reaction
- F, Ruthenium catalyzed domino reaction
- G, Miscellaneous domino reactions

Reviews about domino reactions:

Chem. Rev. **2014**, *114*, 2390; *Chem. Rev.* **2014**, *114*, 8323; *Green Chem.* **2014**, *16*, 2958;
Chem. Soc. Rev. **2014**, *43*, 4368; *Tetrahedron* **2013**, *69*, 7171; *Chem. Soc. Rev.* **2013**, *42*, 4948

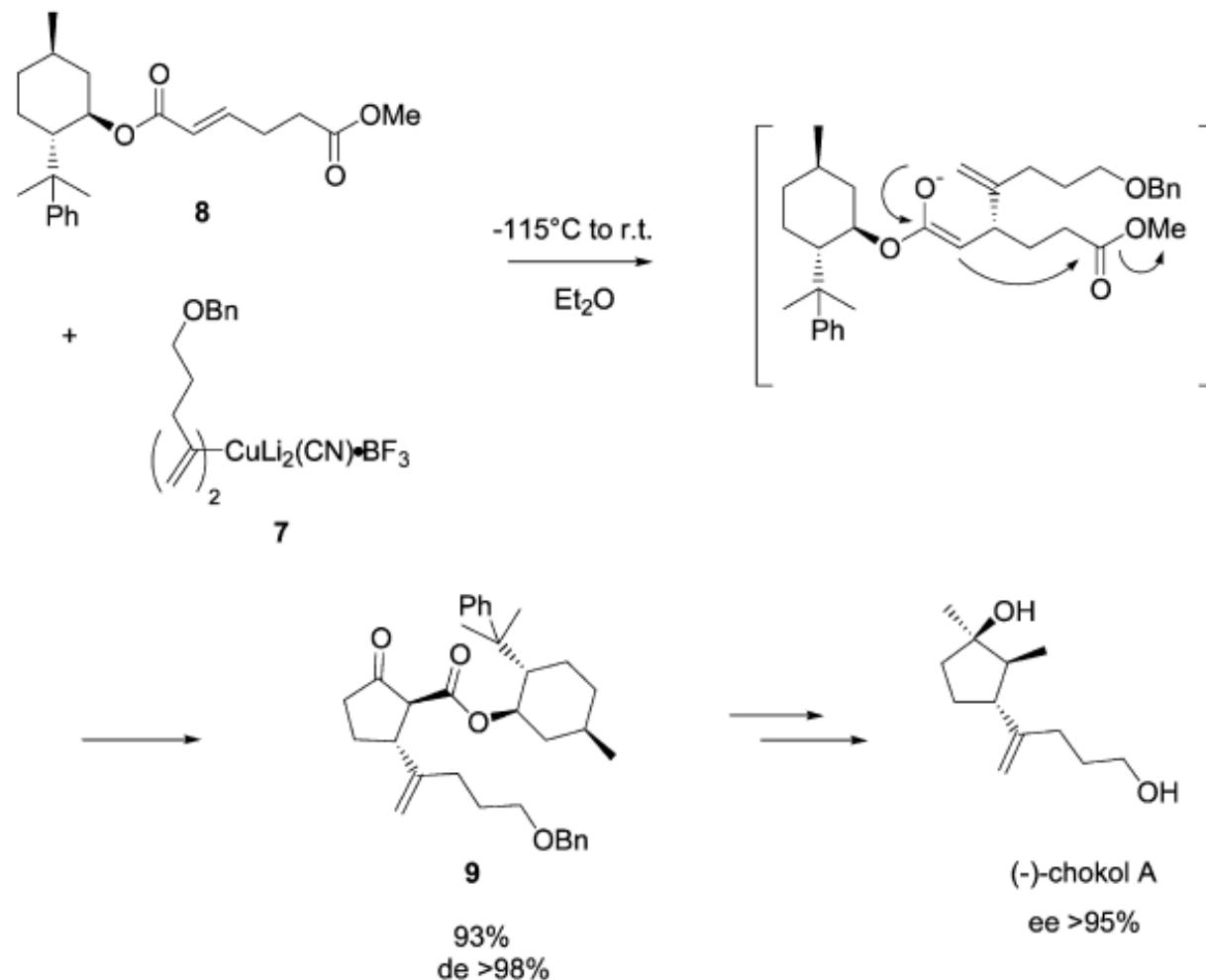
Anionic primary step

Synthesis of Salvinorin A through Domino Michael/Michael Reaction



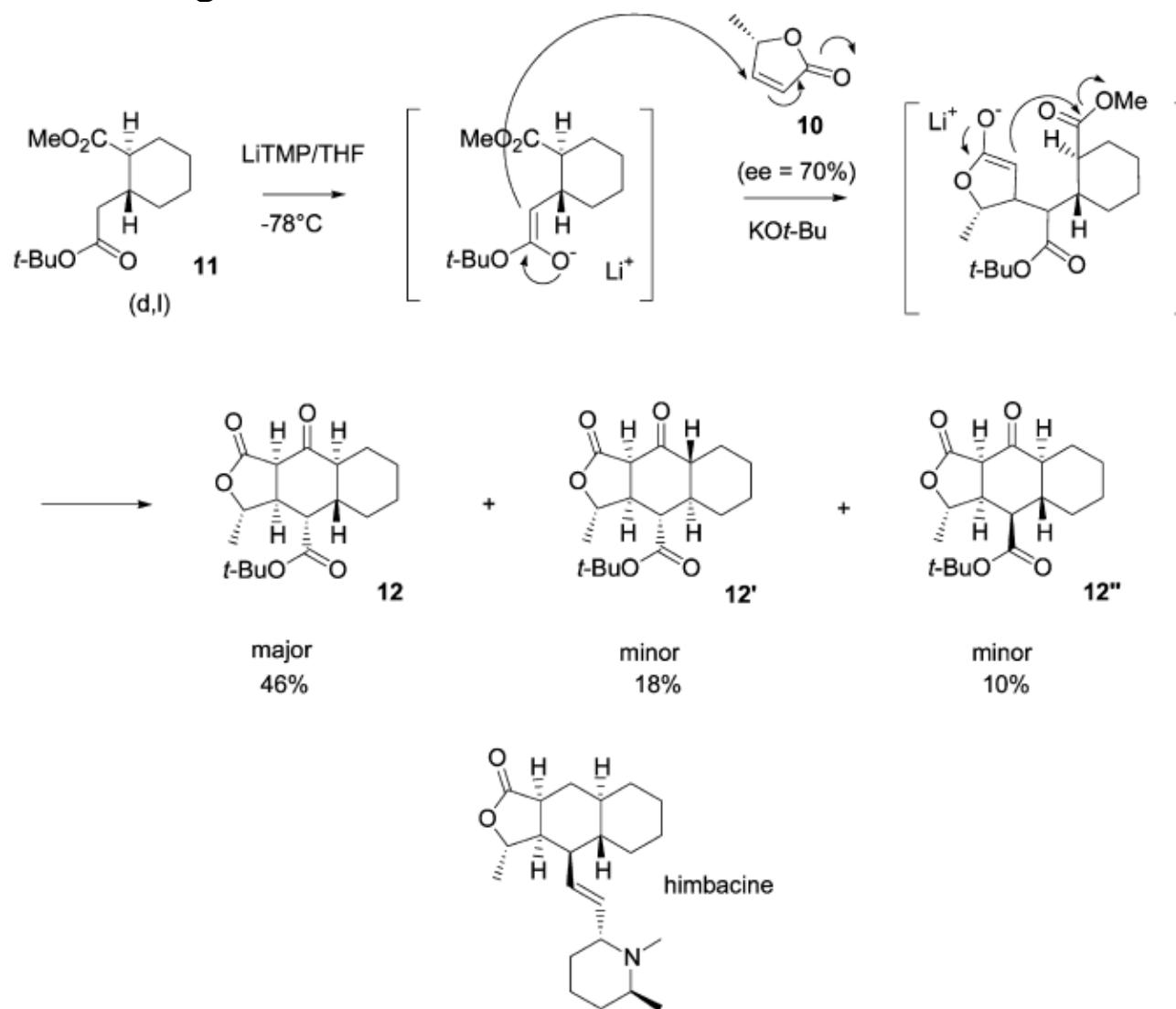
Anionic primary step

Synthesis of (-)-Chokol A through Domino Michael/Dieckmann Reaction



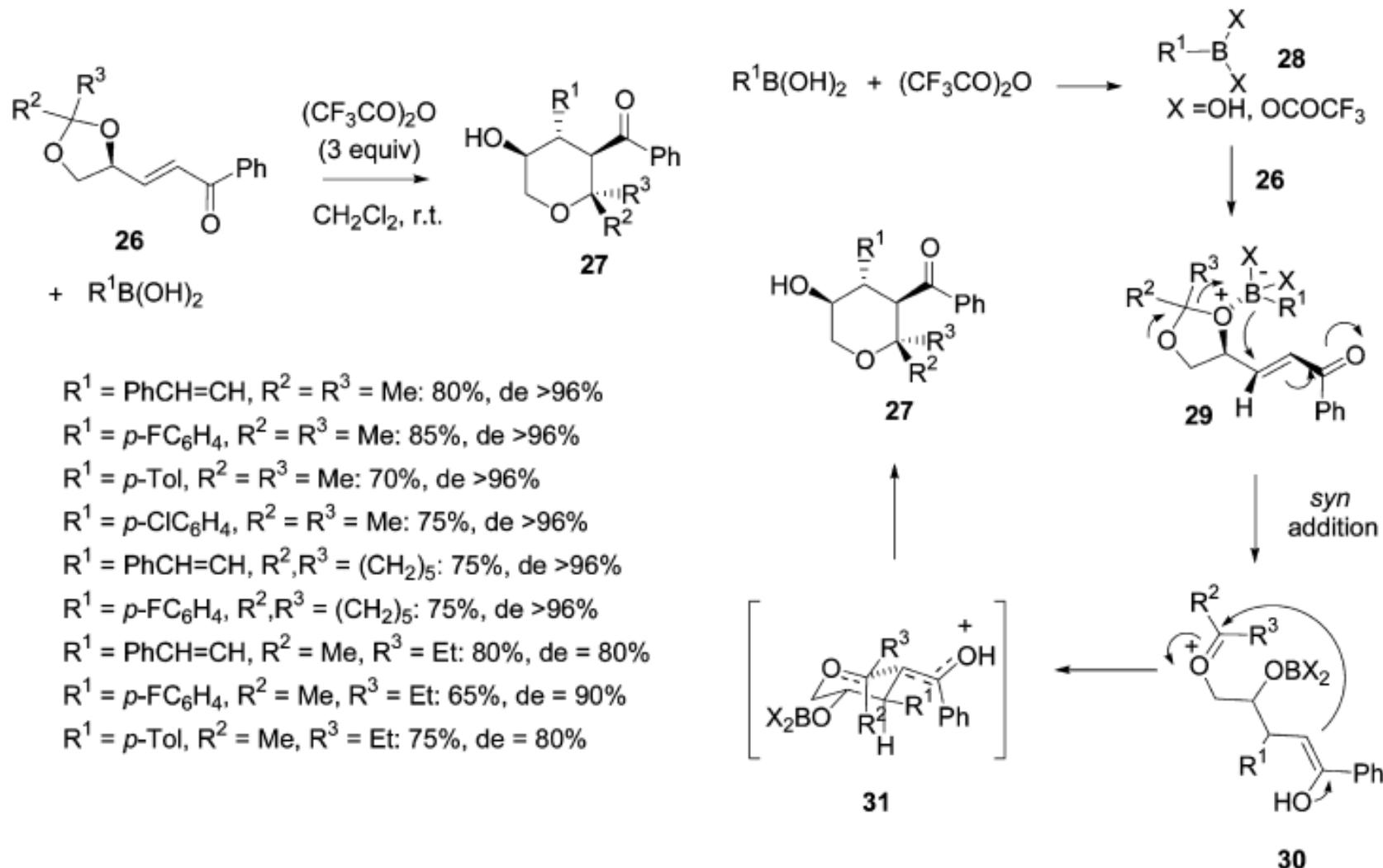
Anionic primary step

Synthesis of Key Intermediate in the Synthesis of Himbacine through Domino Michael/Dieckmann Reaction



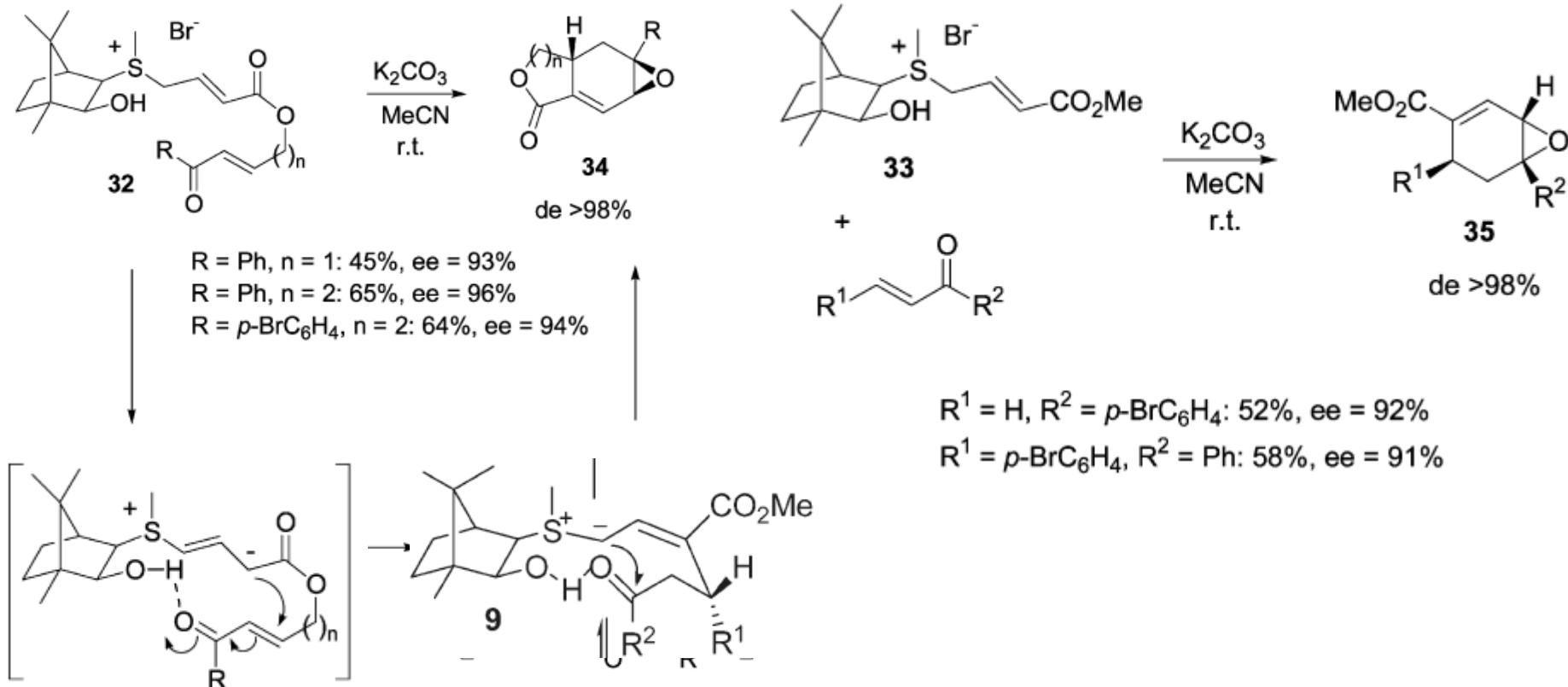
Domino Michael/Acetal Ring-Opening Reaction

proposed mechanism:



Anionic primary step

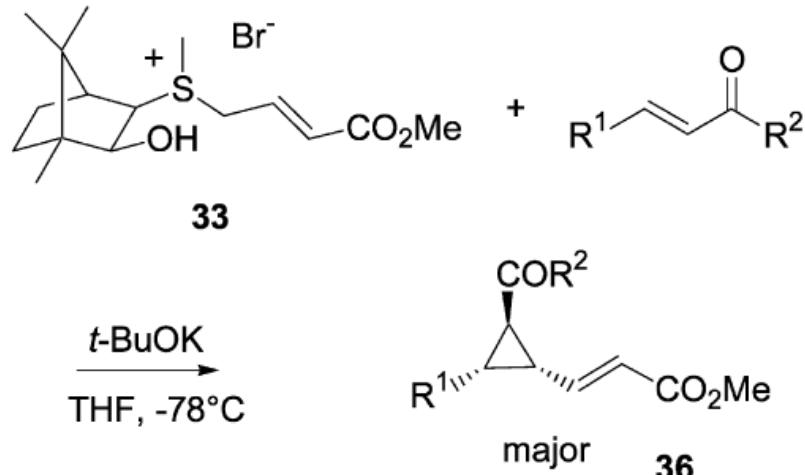
Intra- and Intermolecular Domino Michael/Ylide Epoxidation Reactions



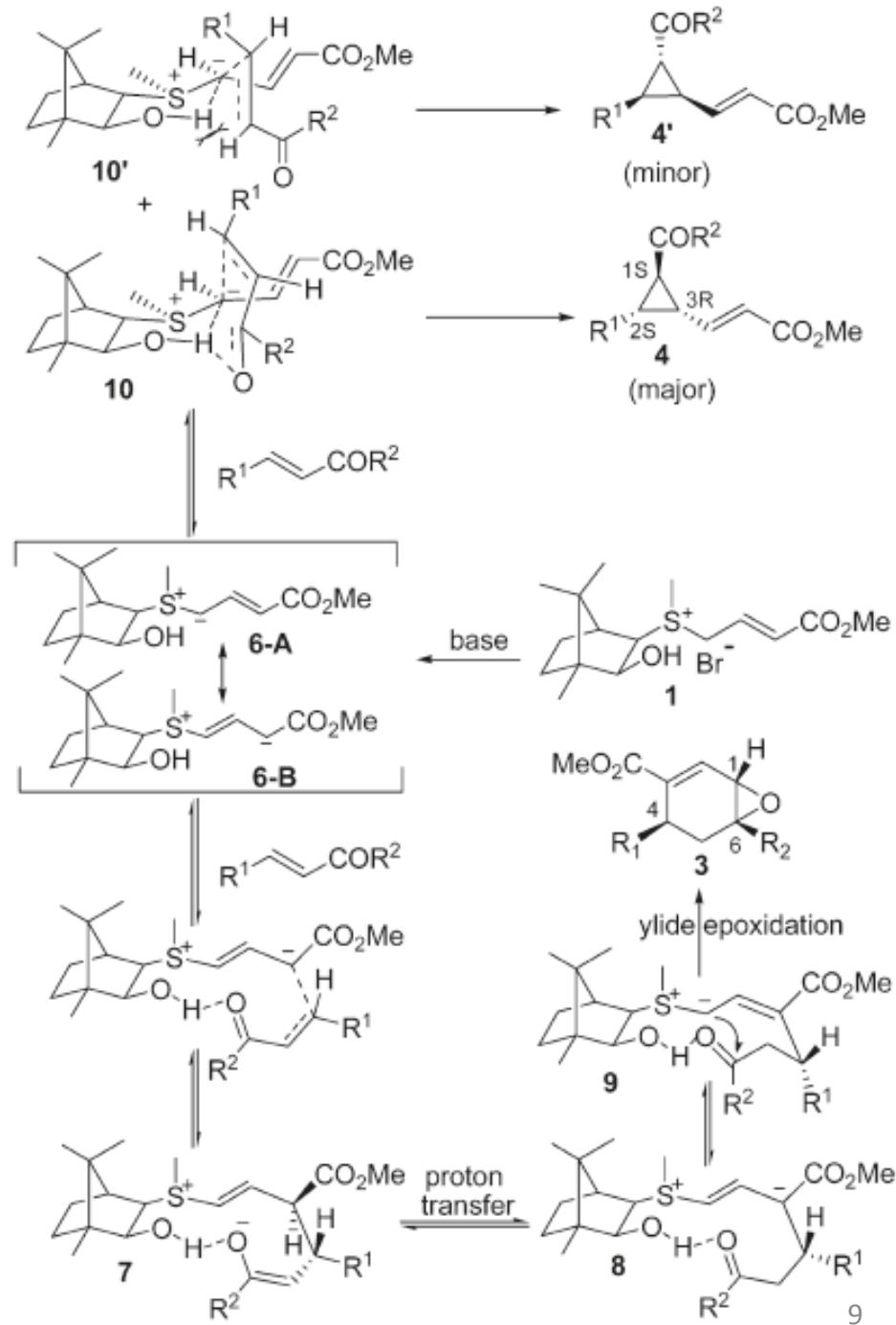
Tang, Y. *J. Am. Chem. Soc.* **2008**, *130*, 5408

Anionic primary step

Domino Michael/Ylide Cyclopropanation Reaction

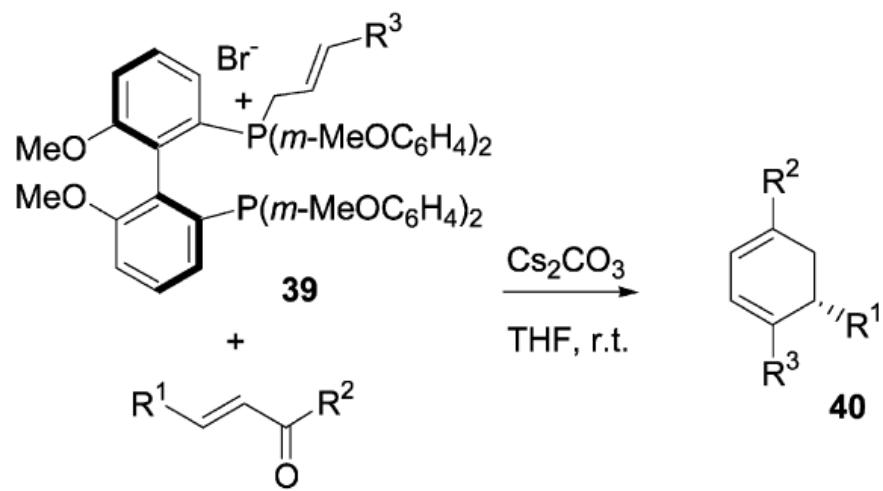


- $R^1 = R^2 = \text{Ph}$: 77%
 $R^1 = p\text{-ClC}_6\text{H}_4$, $R^2 = \text{Ph}$: 57%
 $R^1 = p\text{-BrC}_6\text{H}_4$, $R^2 = \text{Ph}$: 61%
 $R^1 = p\text{-Tol}$, $R^2 = \text{Ph}$: 87%
 $R^1 = p\text{-MeOC}_6\text{H}_4$, $R^2 = \text{Ph}$: 78%
 $R^1 = \text{Ph}$, $R^2 = p\text{-Tol}$: 75%
 $R^1 = \text{Ph}$, $R^2 = p\text{-ClC}_6\text{H}_4$: 56%



Anionic primary step

Domino Michael/Wittig Reaction



$\text{R}^1 = \text{R}^2 = \text{Ph}$, $\text{R}^3 = \text{CO}_2\text{Me}$: 64%, ee = 84%

$\text{R}^1 = \text{R}^2 = \text{Ph}$, $\text{R}^3 = \text{CO}_2t\text{-Bu}$: 55%, ee = 79%

$\text{R}^1 = p\text{-BrC}_6\text{H}_4$, $\text{R}^2 = \text{Ph}$, $\text{R}^3 = \text{CO}_2\text{Me}$: 73%, ee = 80%

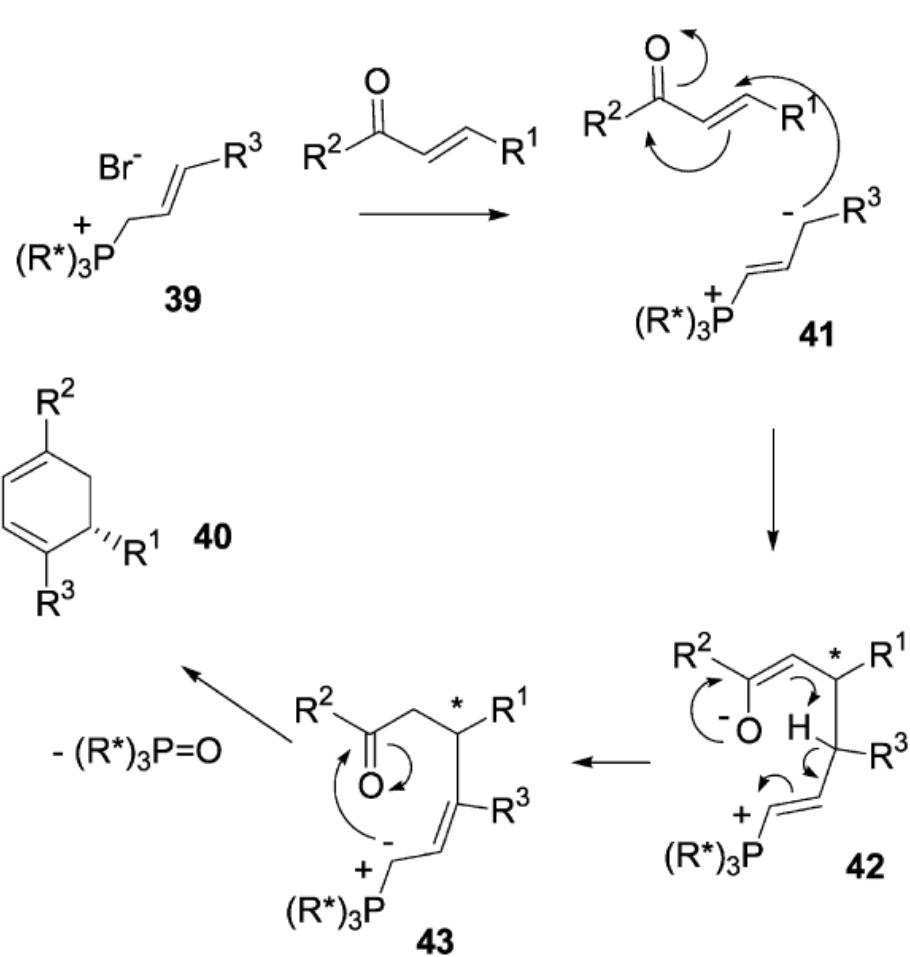
$\text{R}^1 = o\text{-BrC}_6\text{H}_4$, $\text{R}^2 = \text{Ph}$, $\text{R}^3 = \text{CO}_2\text{Me}$: 60%, ee = 80%

$\text{R}^1 = p\text{-MeOC}_6\text{H}_4$, $\text{R}^2 = \text{Ph}$, $\text{R}^3 = \text{CO}_2\text{Me}$: 65%, ee = 90%

$\text{R}^1 = \text{Me}$, $\text{R}^2 = \text{Ph}$, $\text{R}^3 = \text{CO}_2\text{Me}$: 53%, ee = 78%

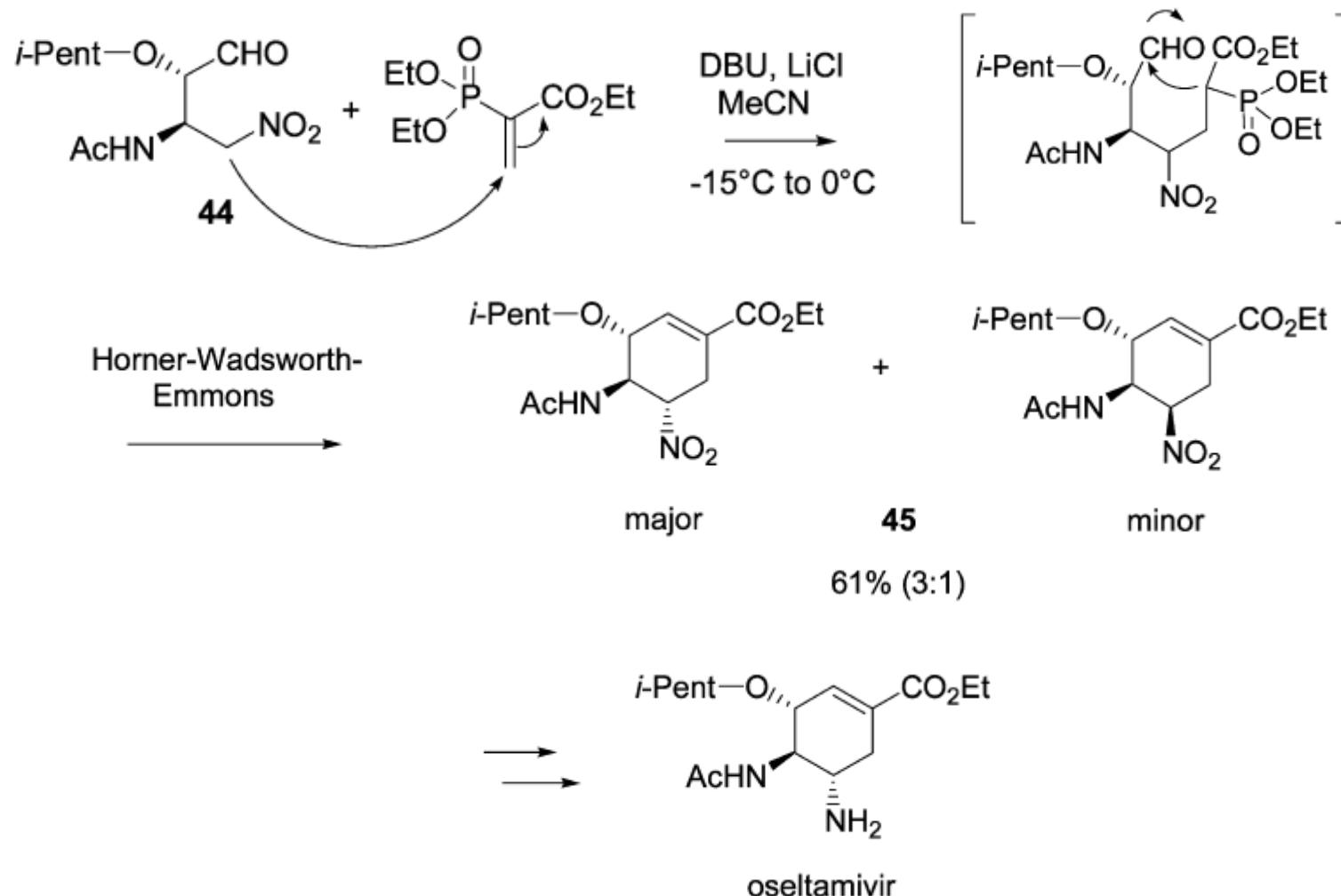
$\text{R}^1 = \text{Me}$, $\text{R}^2 = \text{H}$, $\text{R}^3 = \text{CO}_2\text{Me}$: 67%, ee = 25%

proposed mechanism:



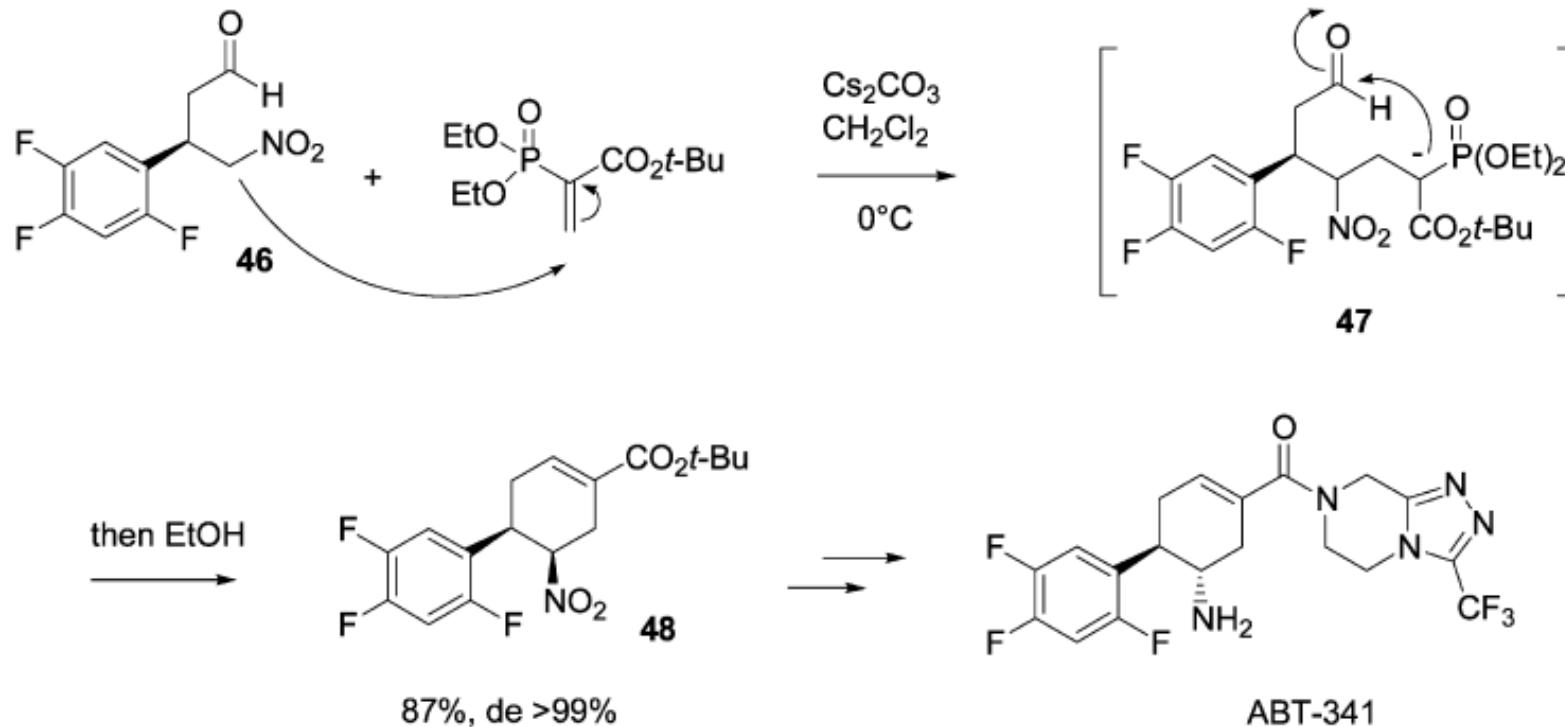
Anionic primary step

Synthesis of Oseltamivir through Domino Michael/Horner-Wadsworth-Emmons Reaction



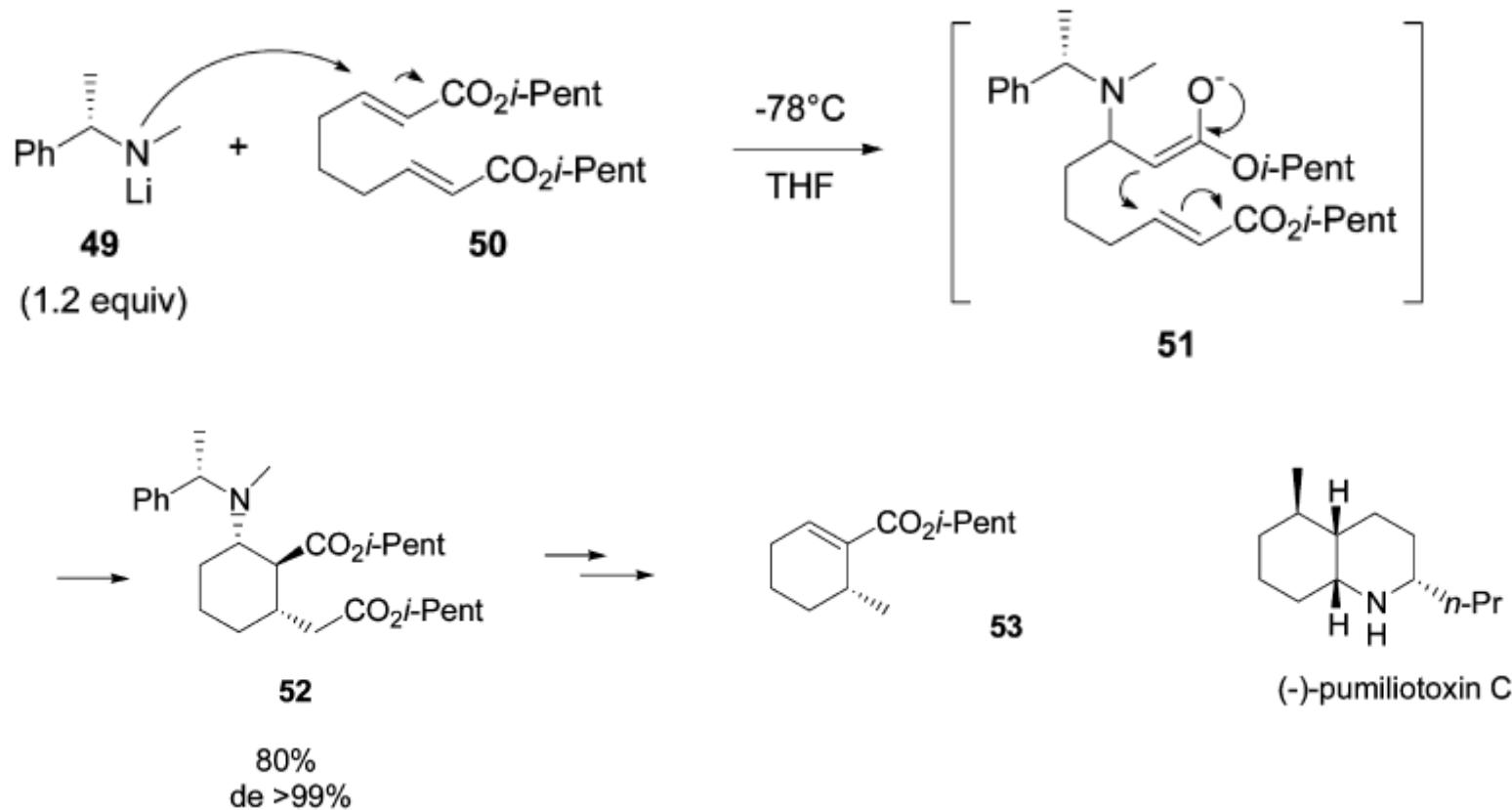
Anionic primary step

Synthesis of ABT-341 through Domino Michael/Horner-Wadsworth-Emmons Reaction



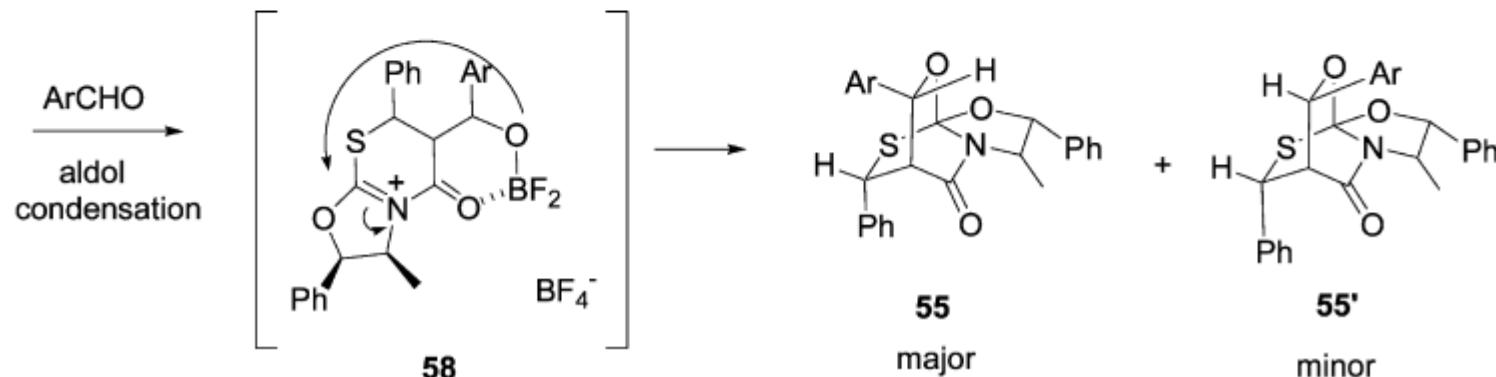
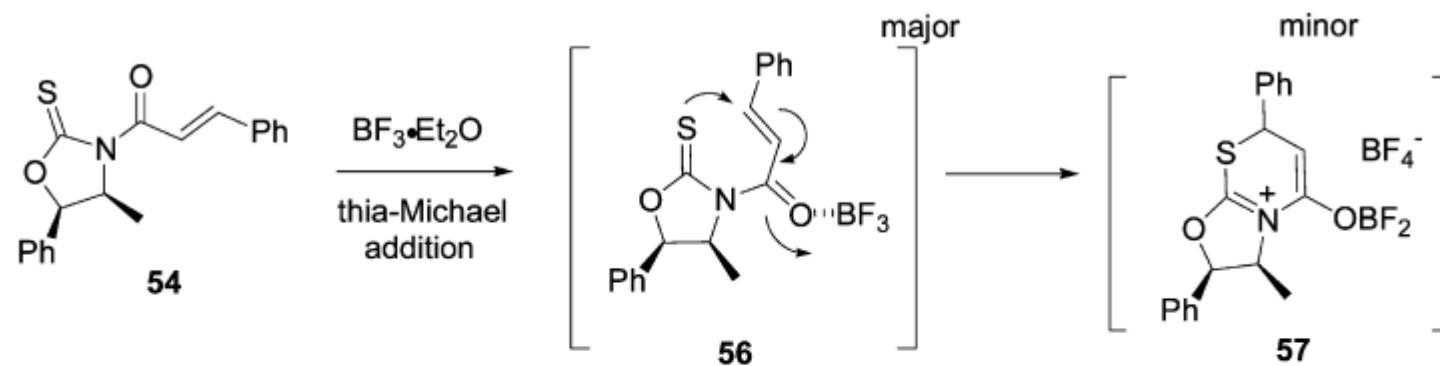
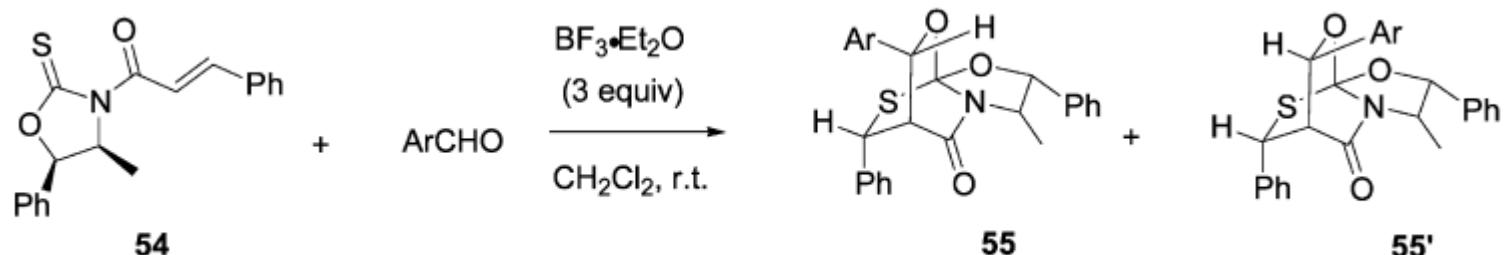
Anionic primary step

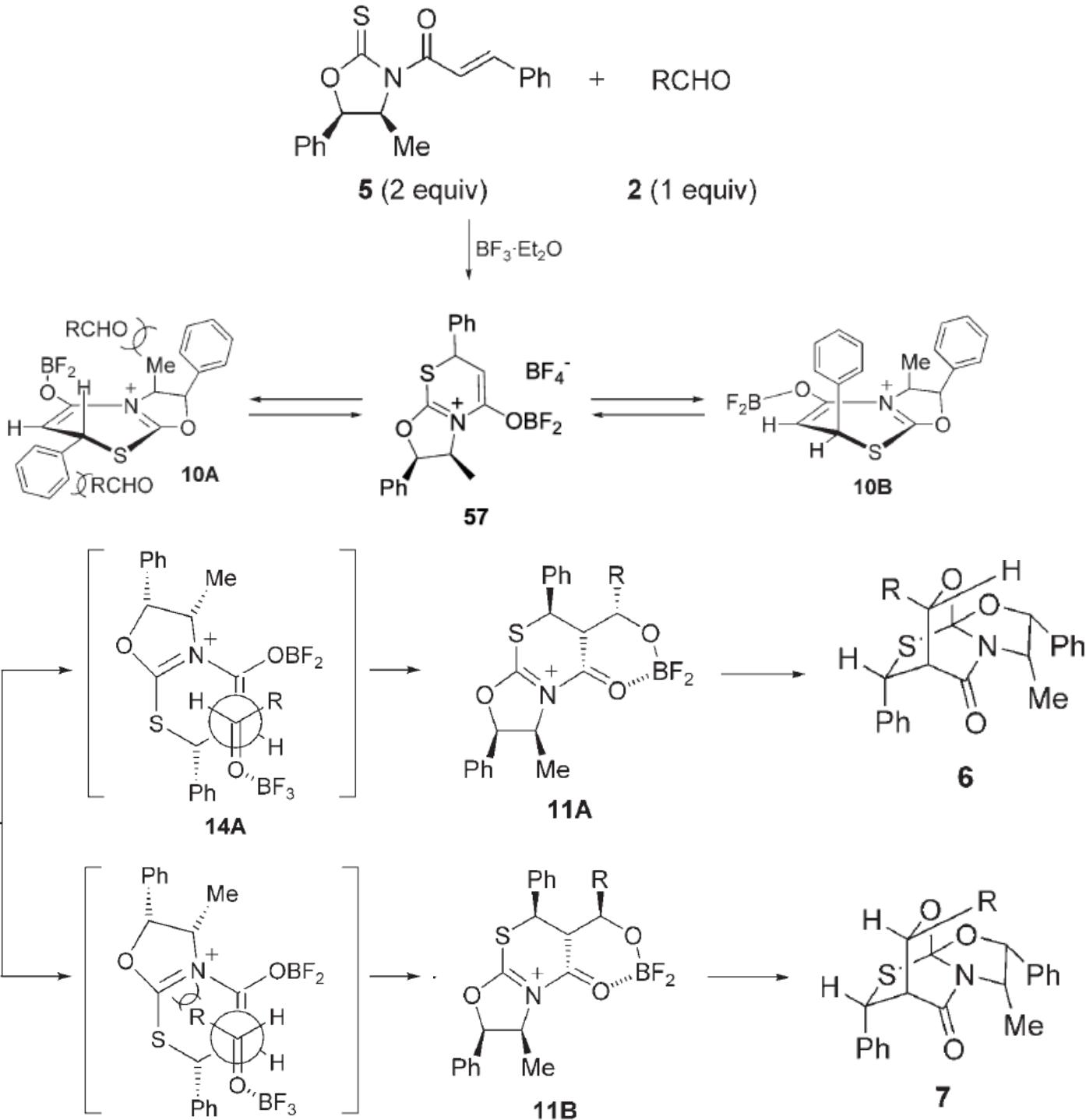
Synthesis of Precursor of (-)-Pumiliotoxin C through Domino Aza-Michael/Michael Reaction



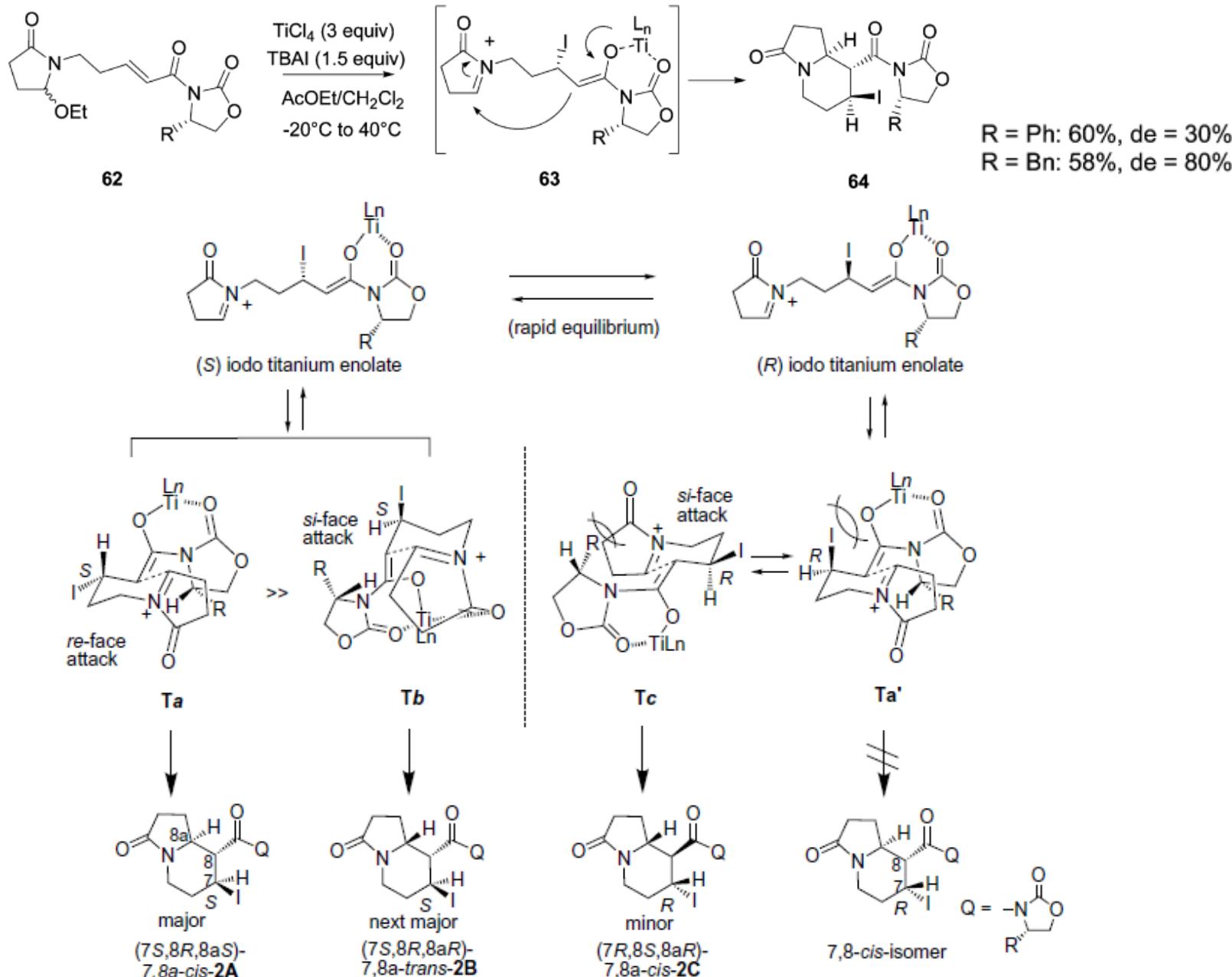
Anionic primary step

Domino Thia-Michael/Aldol Reaction of Chiral 3-Cinnamoyloxazolidine-2-thione with Aromatic Aldehydes



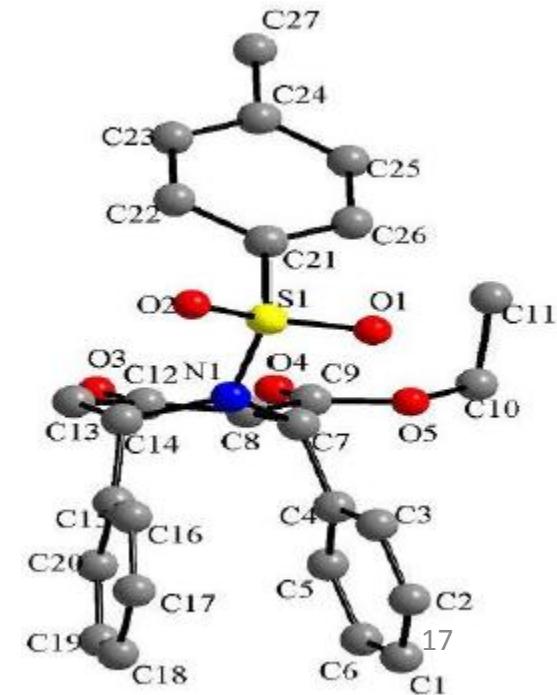
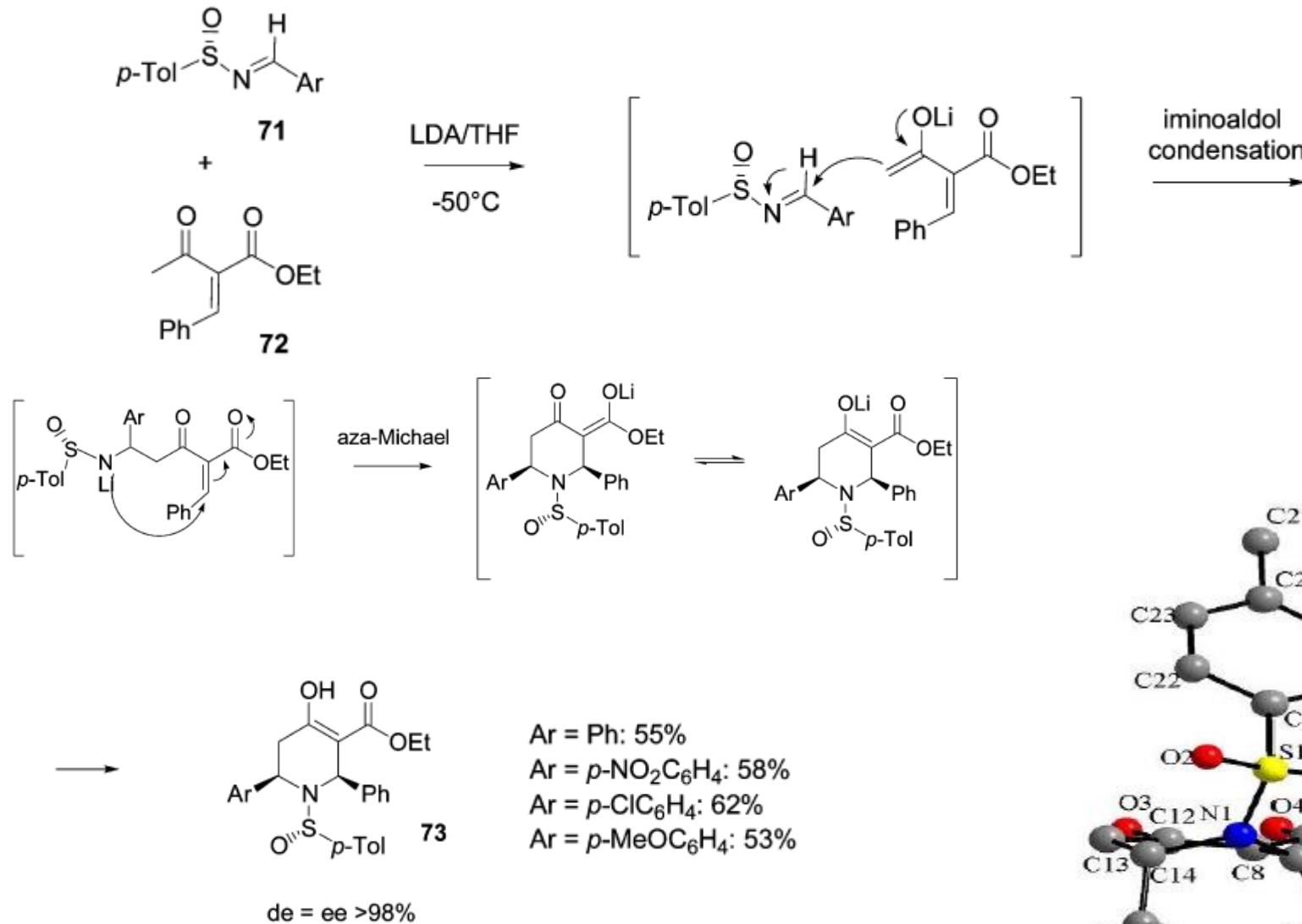


Intramolecular Domino Hetero-Michael/Mannich-type Reaction

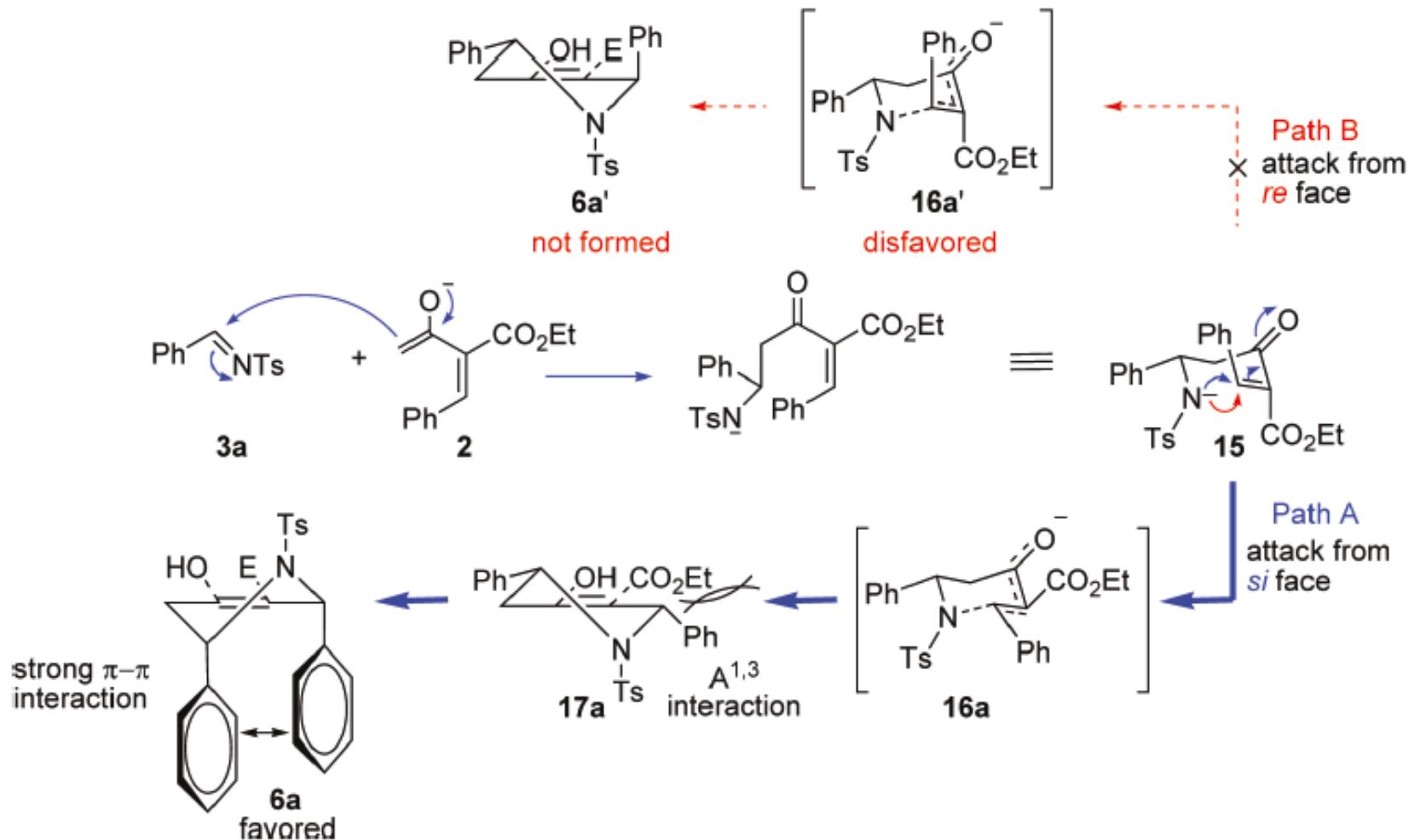


Anionic primary step

Domino Imino-Aldol/Aza-Michael Reaction

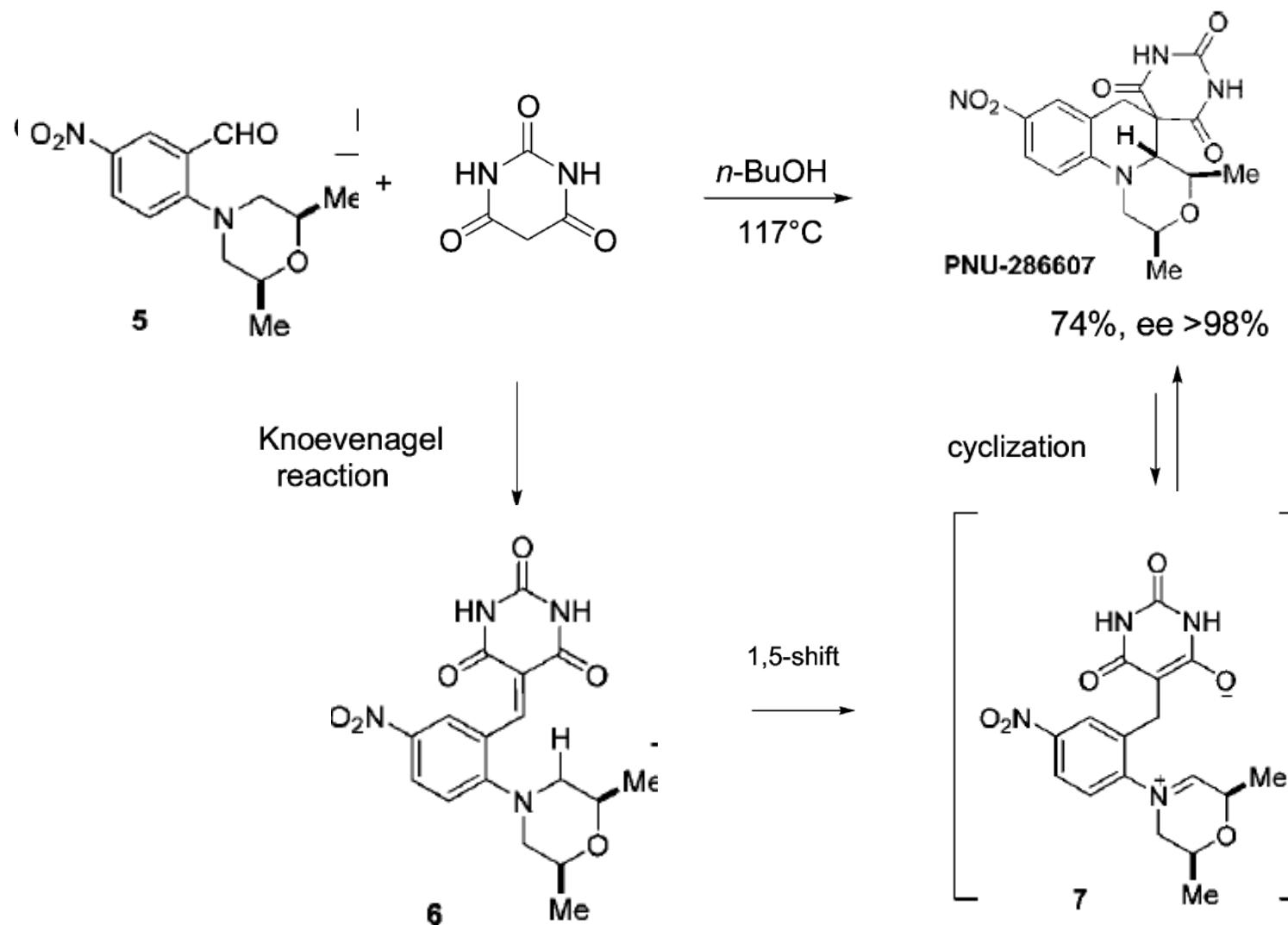


Domino Imino-Aldol/Aza-Michael Reaction

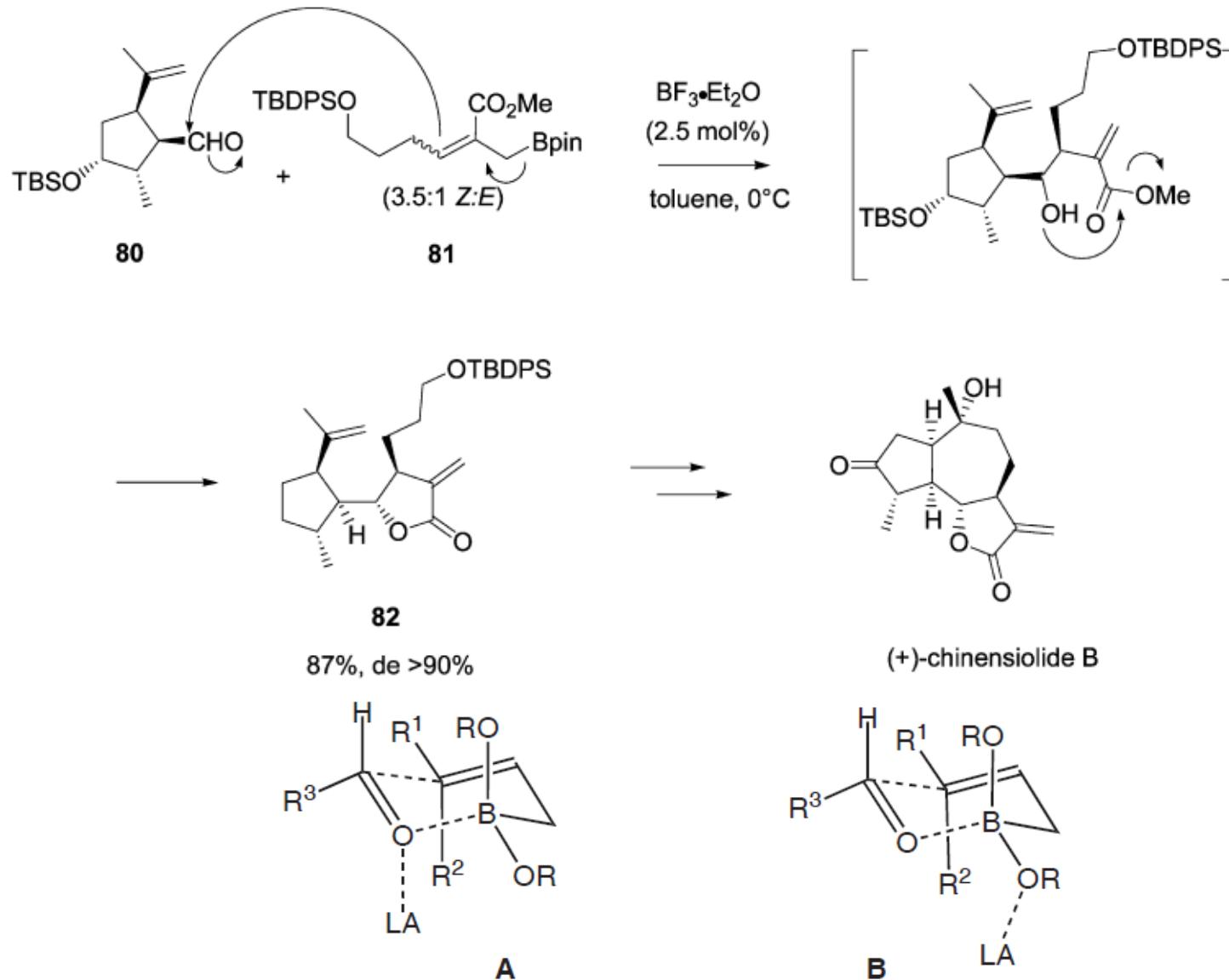


Anionic primary step

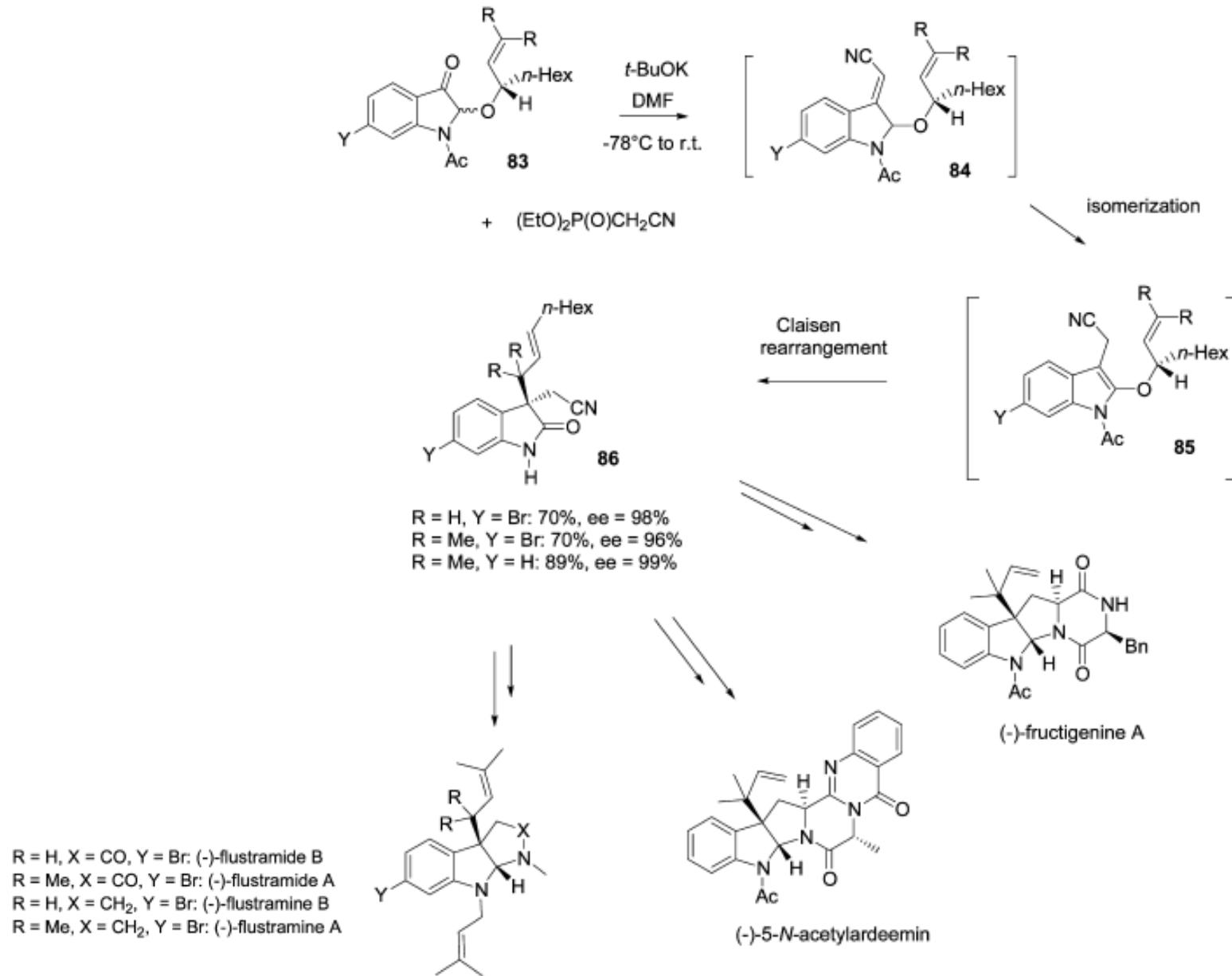
Synthesis of (-)-PNU-286607 through Domino Knoevenagel/[1,5]-Hydrogen Shift/Cyclization Reaction



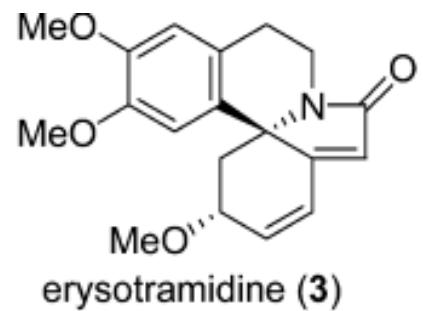
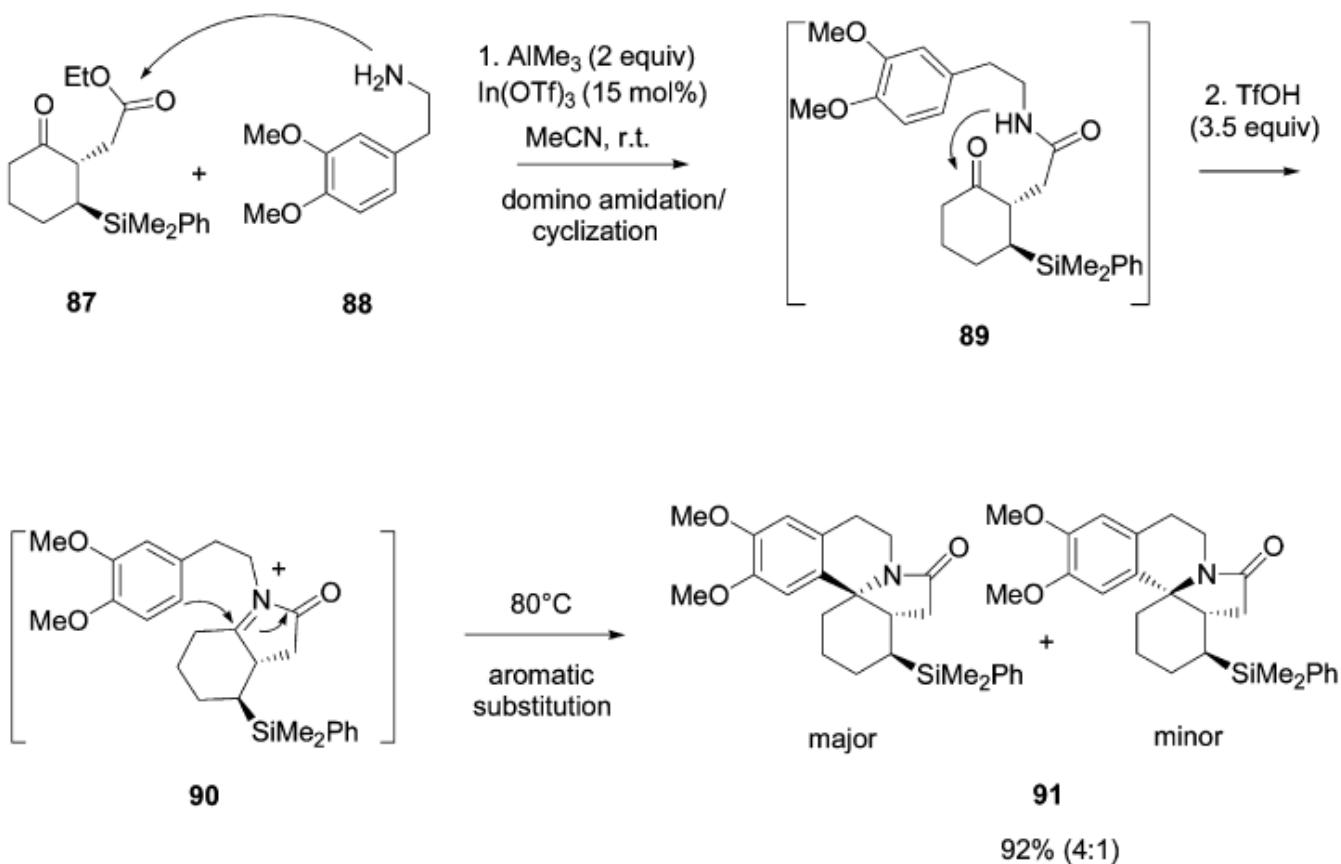
Synthesis of (+)-Chinensiolide B through Domino Allylboration/Lactonization Reaction



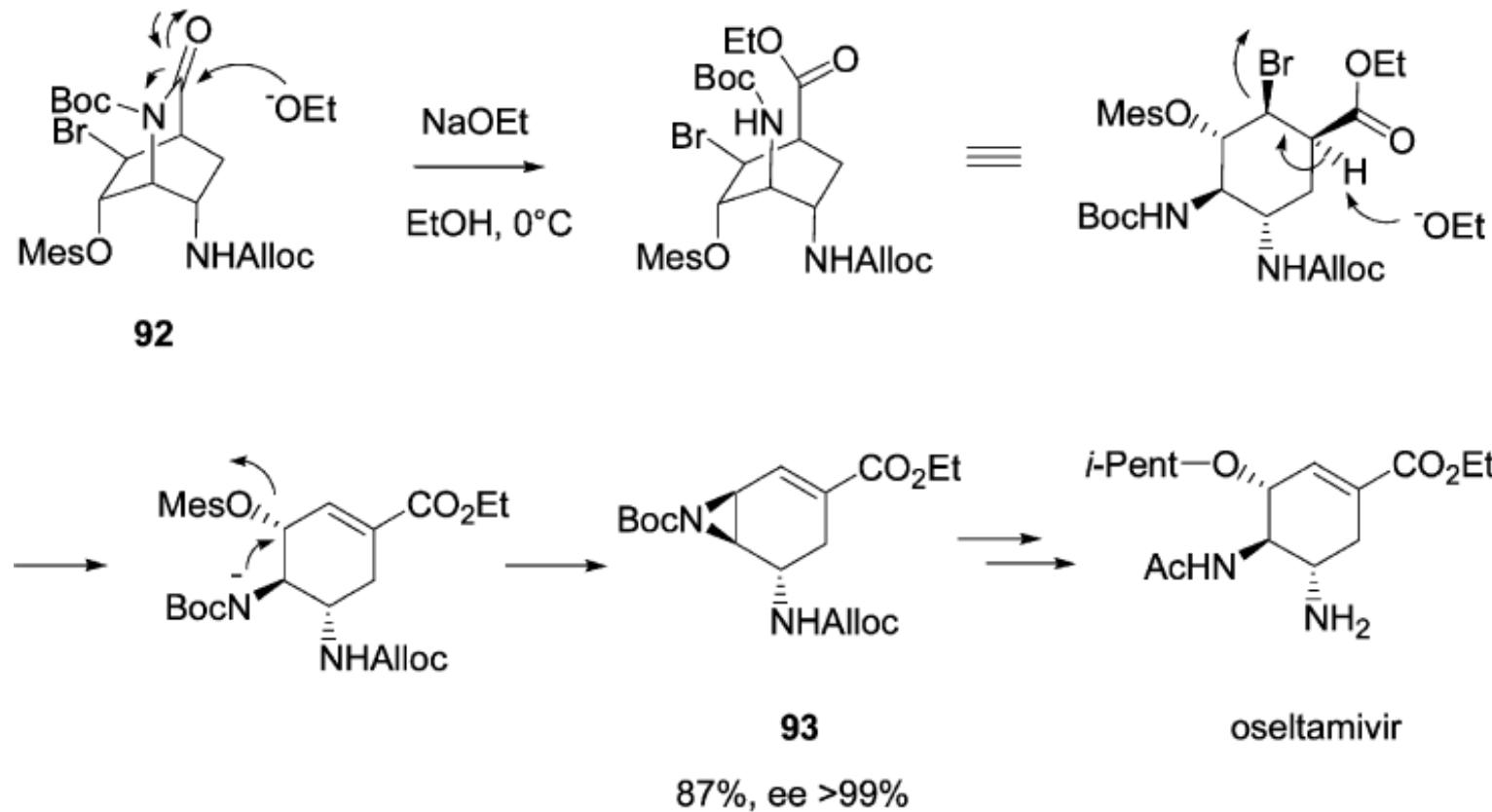
Synthesis of Alkaloids through Domino Horner-Wadsworth-Emmons/Isomerization/Claisen Rearrangement Reaction



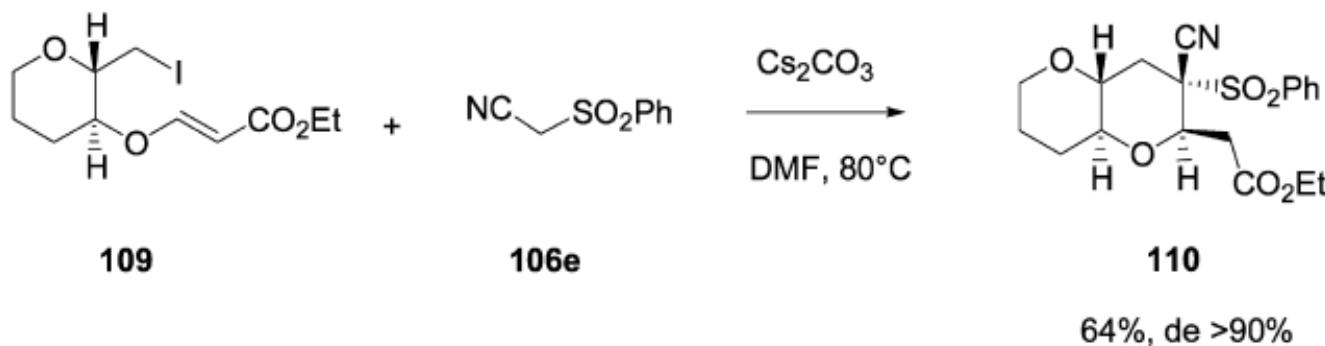
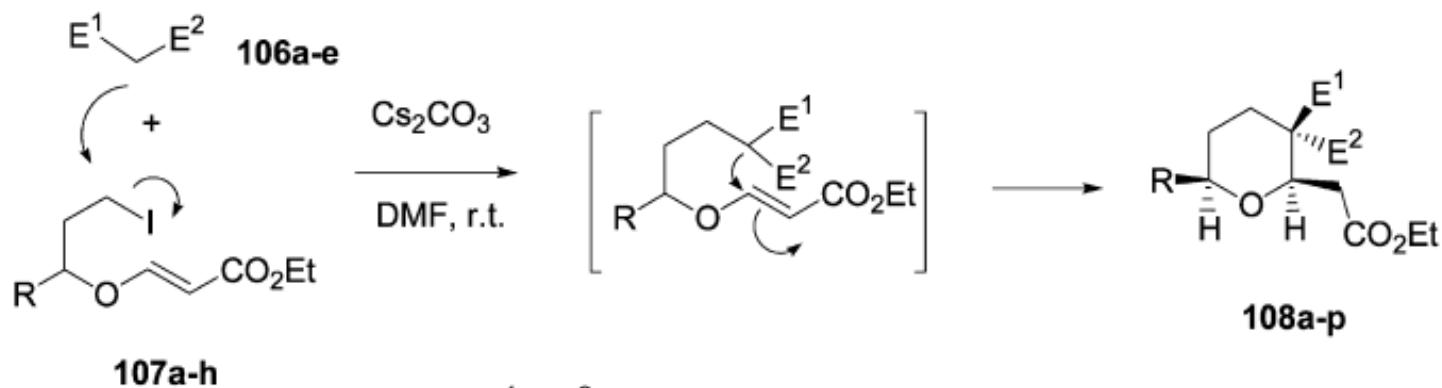
Domino Amidation/Cyclization Reaction Followed by Intramolecular Electrophilic Aromatic Substitution



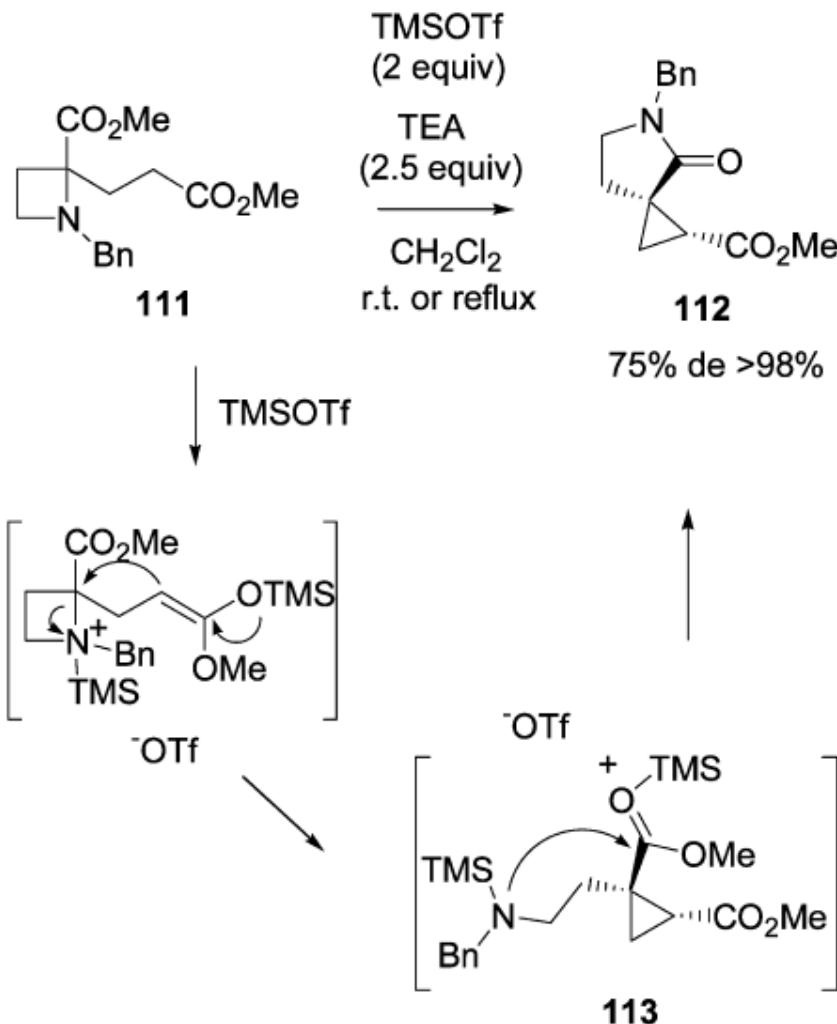
Domino Ethanolysis/Dehydrobromination/Aziridination Reaction



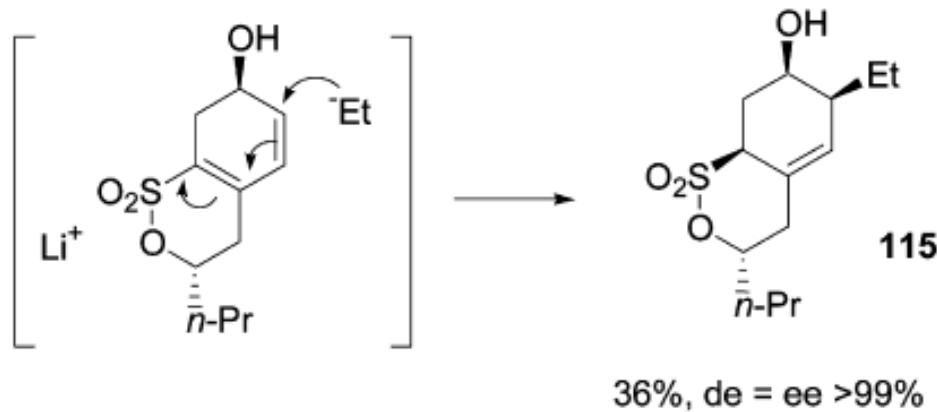
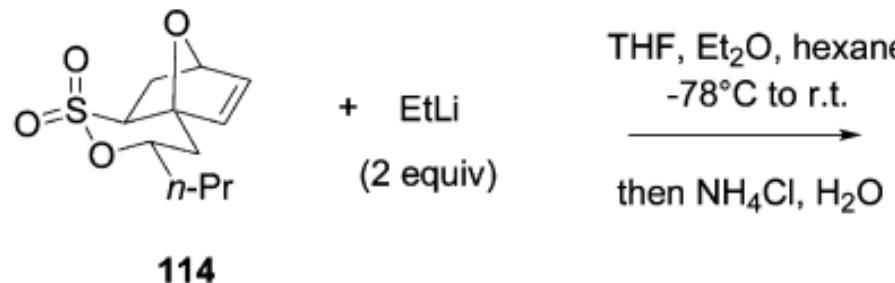
Domino SN₂/Michael Reactions



Domino Intramolecular Azetidine RingOpening/Closing Reaction

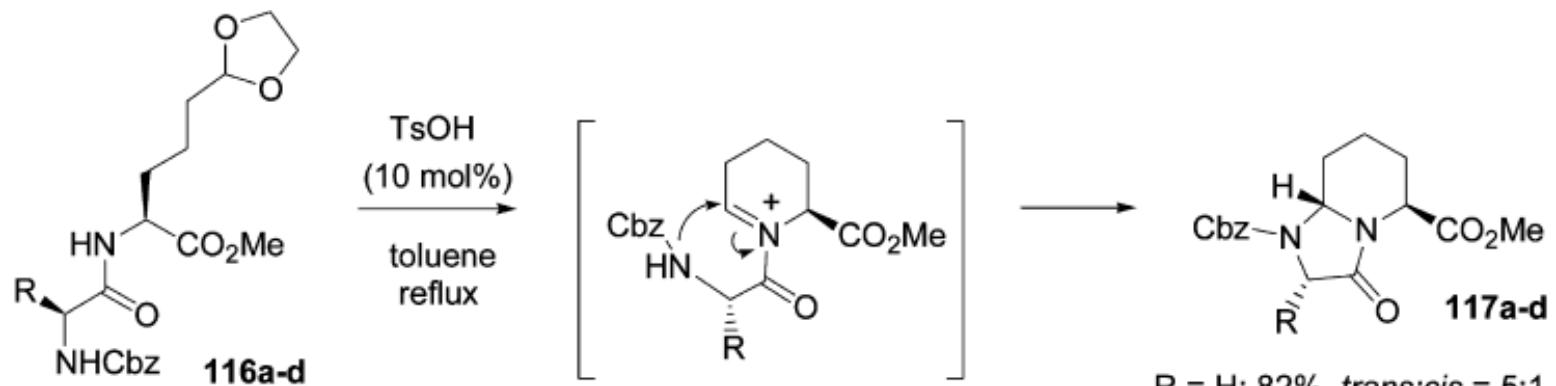


Domino Elimination/1,6-Addition Reaction

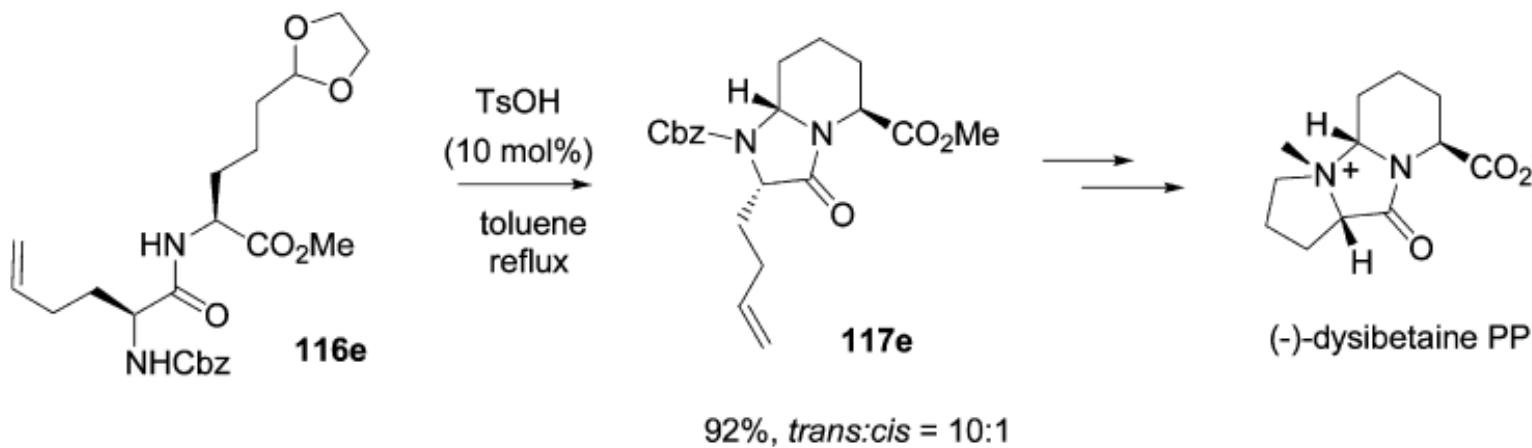


Cationic sequence

Synthesis of (-)-Dysibetaine PP through Domino Cyclization Reaction

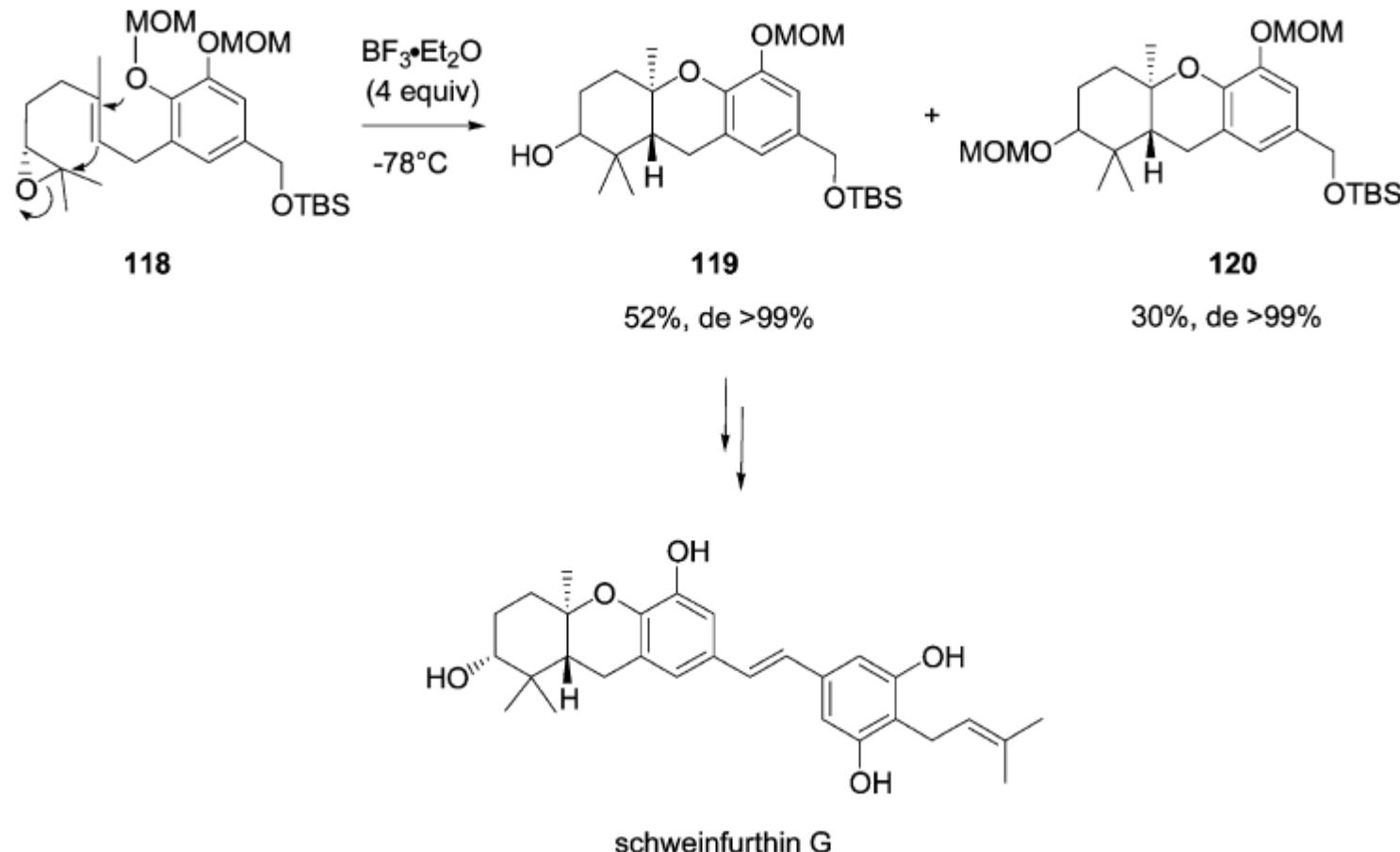


R = H: 82%, *trans:cis* = 5:1
 R = *i*-Pr: 76%, *trans:cis* = 9:1
 R = Bn: 87%, *trans:cis* = 13:1
 R = Allyl: 78%, *trans:cis* = 15:1



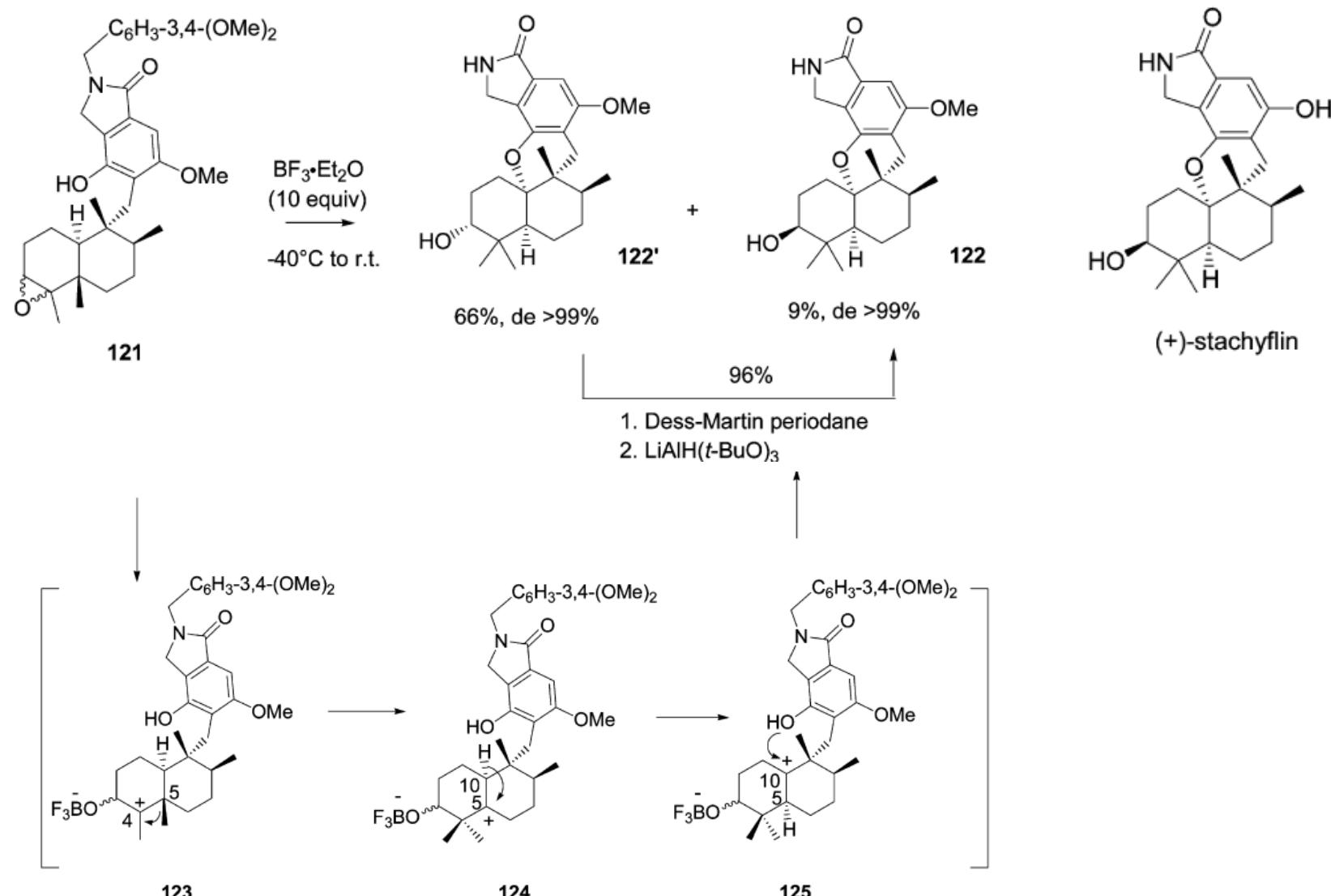
Cationic sequence

Synthesis of Schweinfurthin G through Domino Epoxide-Opening/Cyclization Reaction



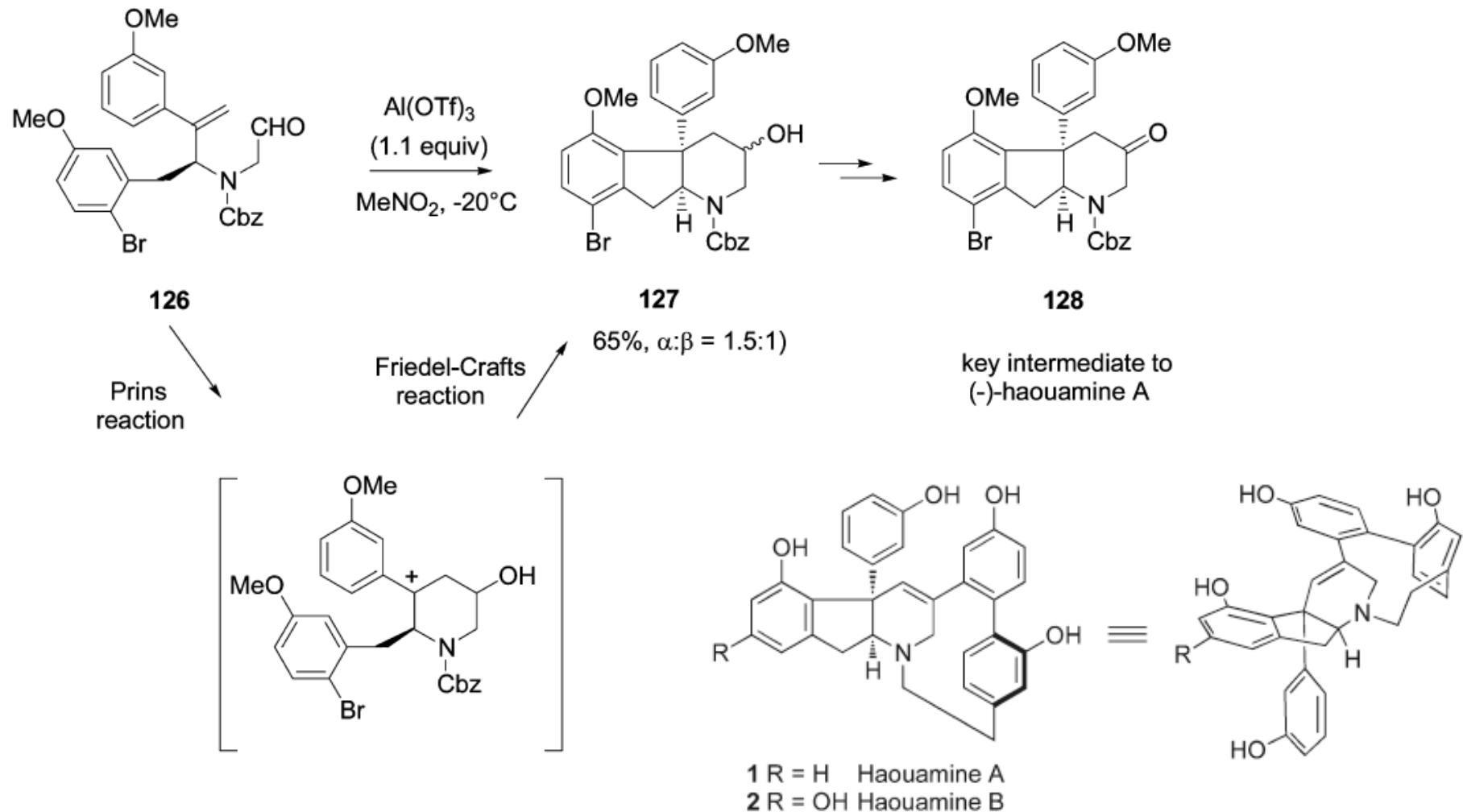
Cationic sequence

Synthesis of (+)-Stachyflin through Domino Epoxide-Opening/Rearrangement/Cyclization Reaction



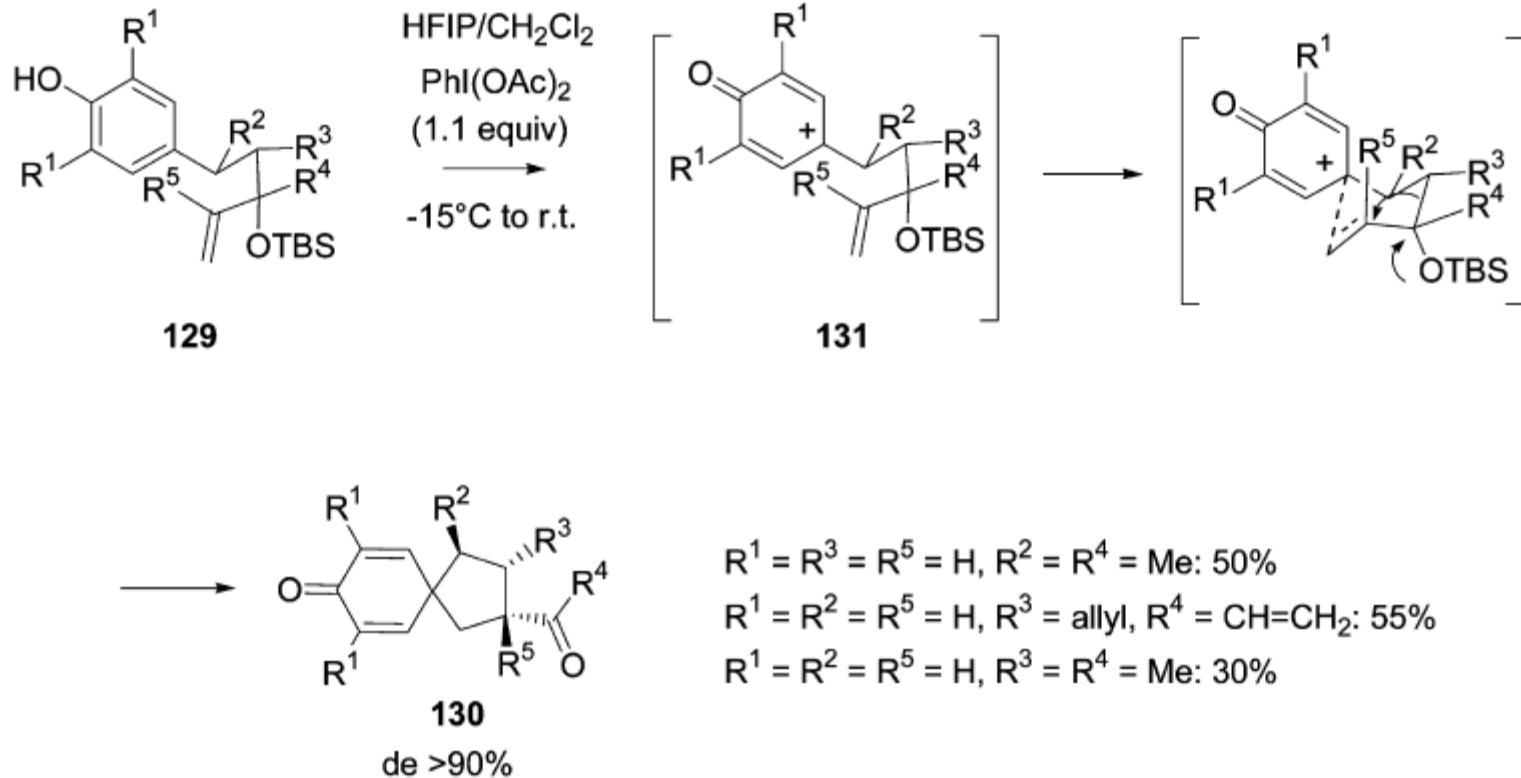
Cationic sequence

Domino Prins/Friedel-Crafts Reaction



Cationic sequence

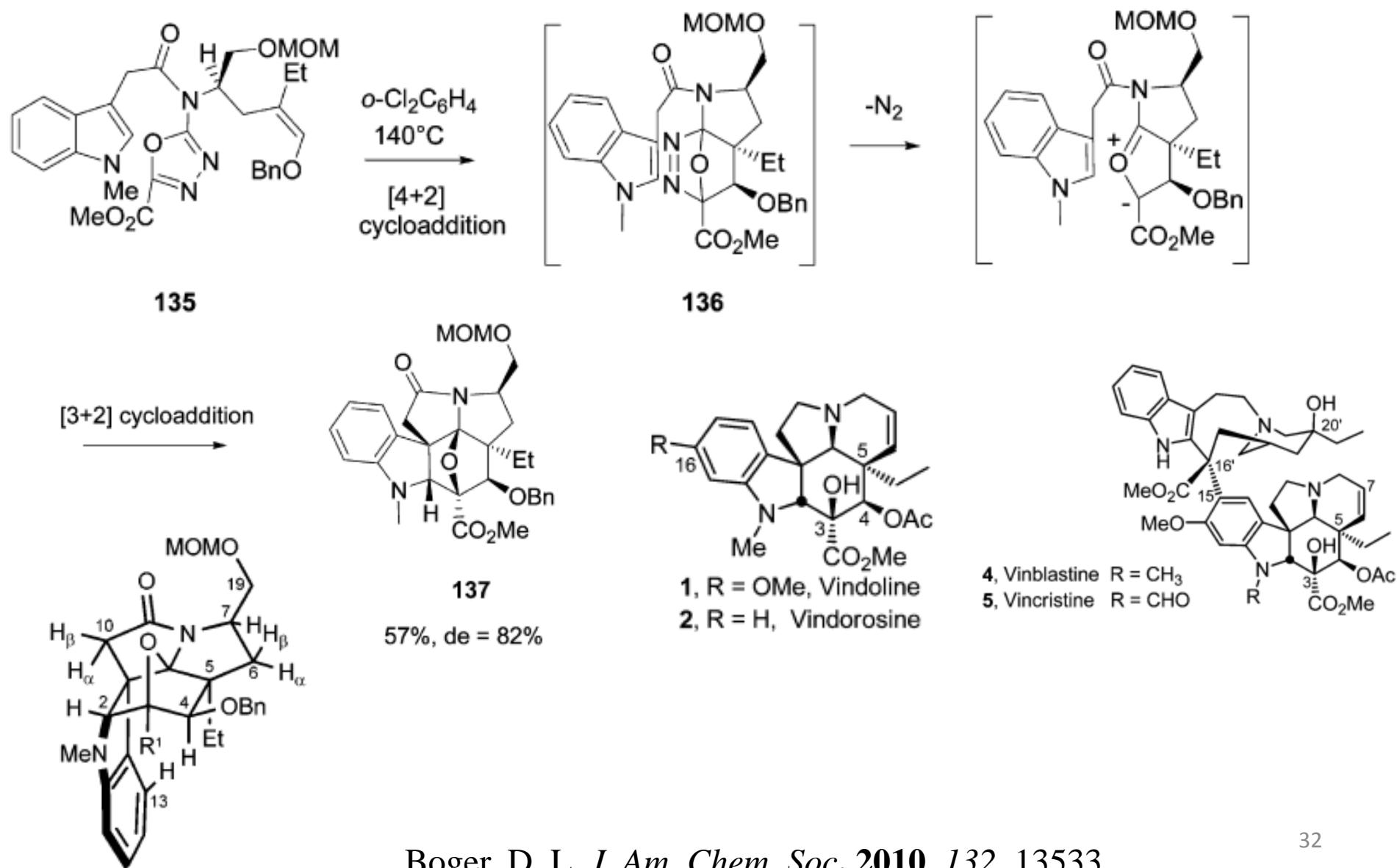
Oxidative Domino Prins/Pinacol Reaction



Canesi, S. *J. Org. Chem.* **2011**, 76, 9460

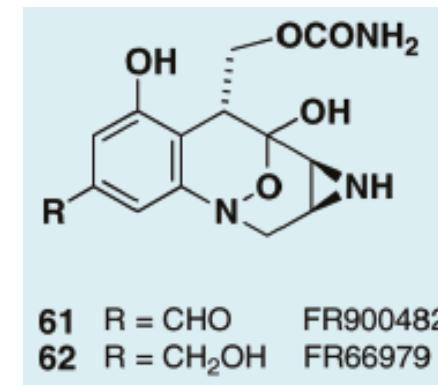
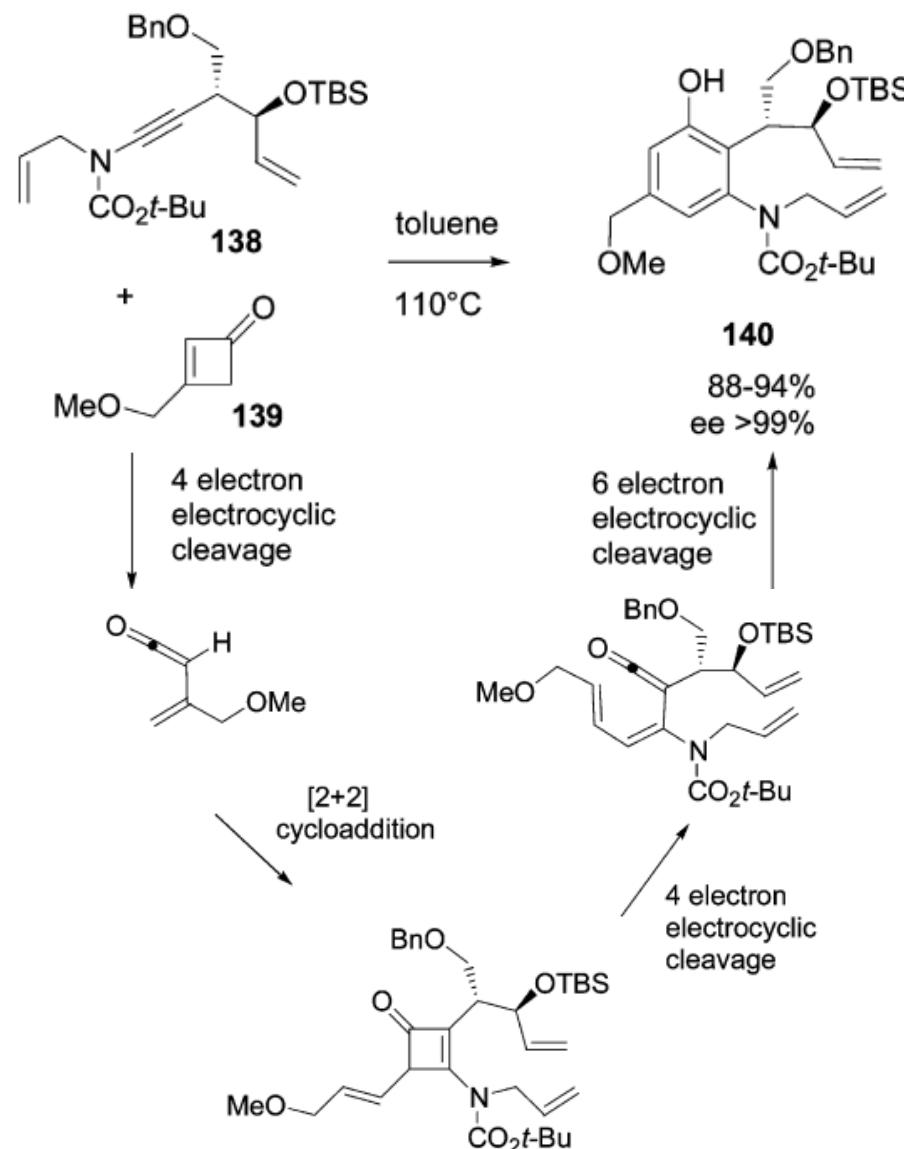
Initiated by a pericyclic primary step

Domino Intramolecular Diels-Alder Cycloaddition/1,3-Dipolar Cycloaddition Reaction



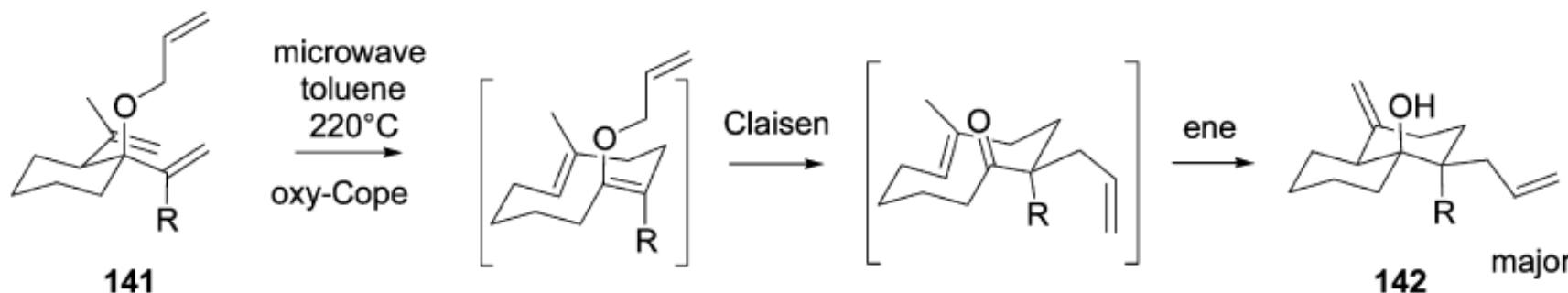
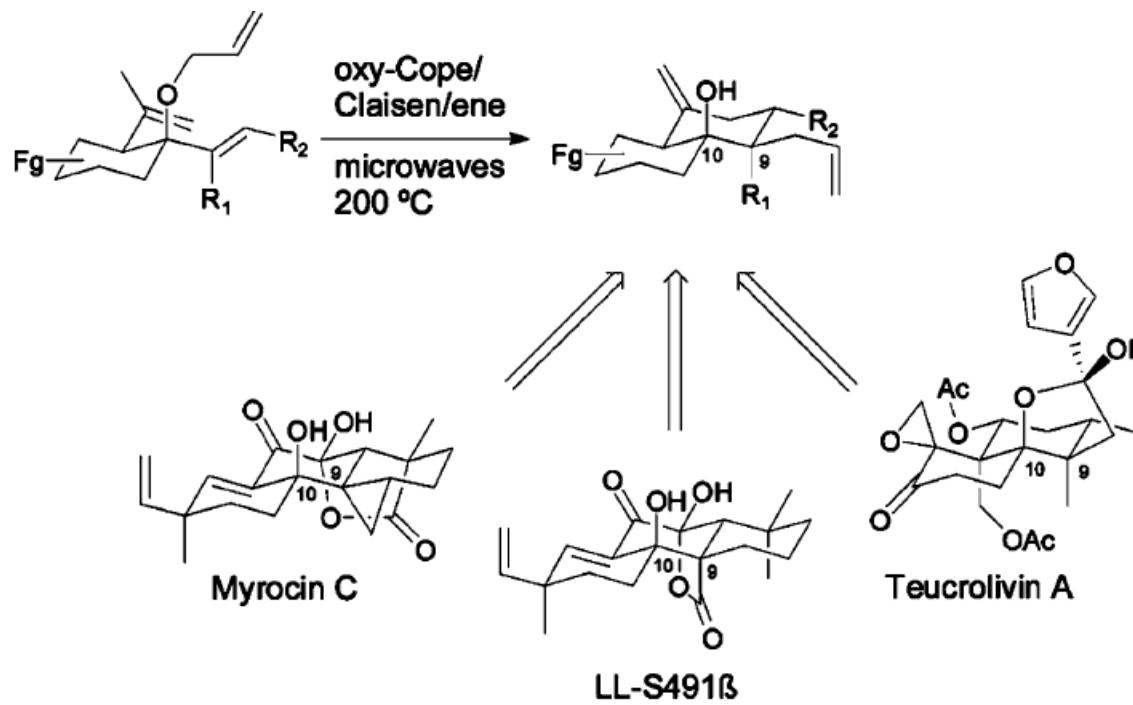
Initiated by a pericyclic primary step

Domino Intramolecular Ynamide Benzannulation Reaction



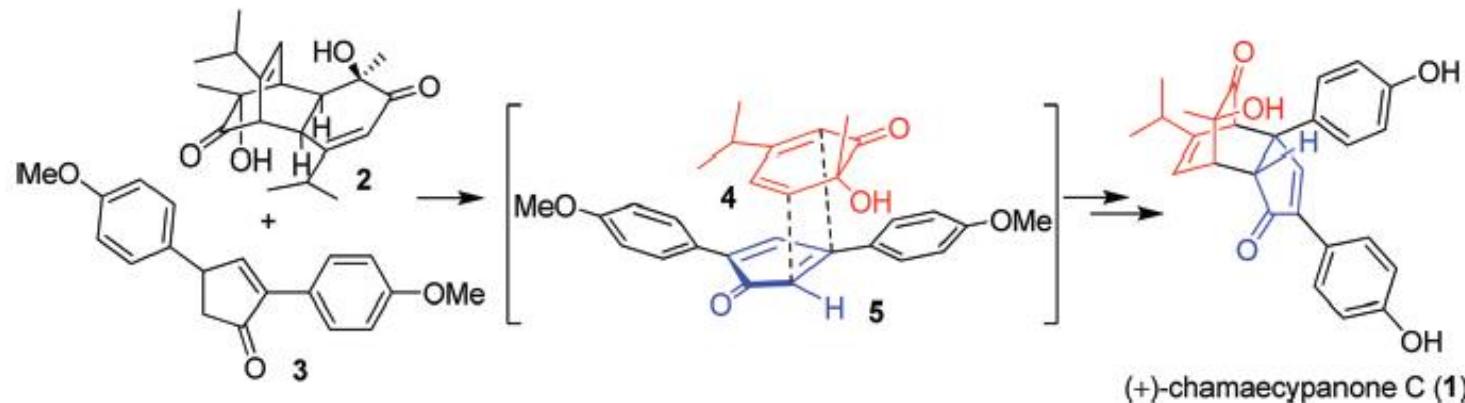
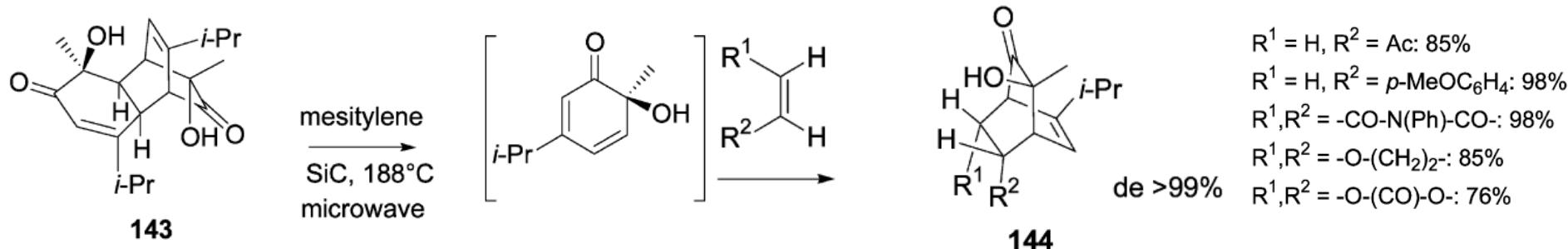
Initiated by a pericyclic primary step

Domino Oxy-Cope/Claisen/Ene Reaction

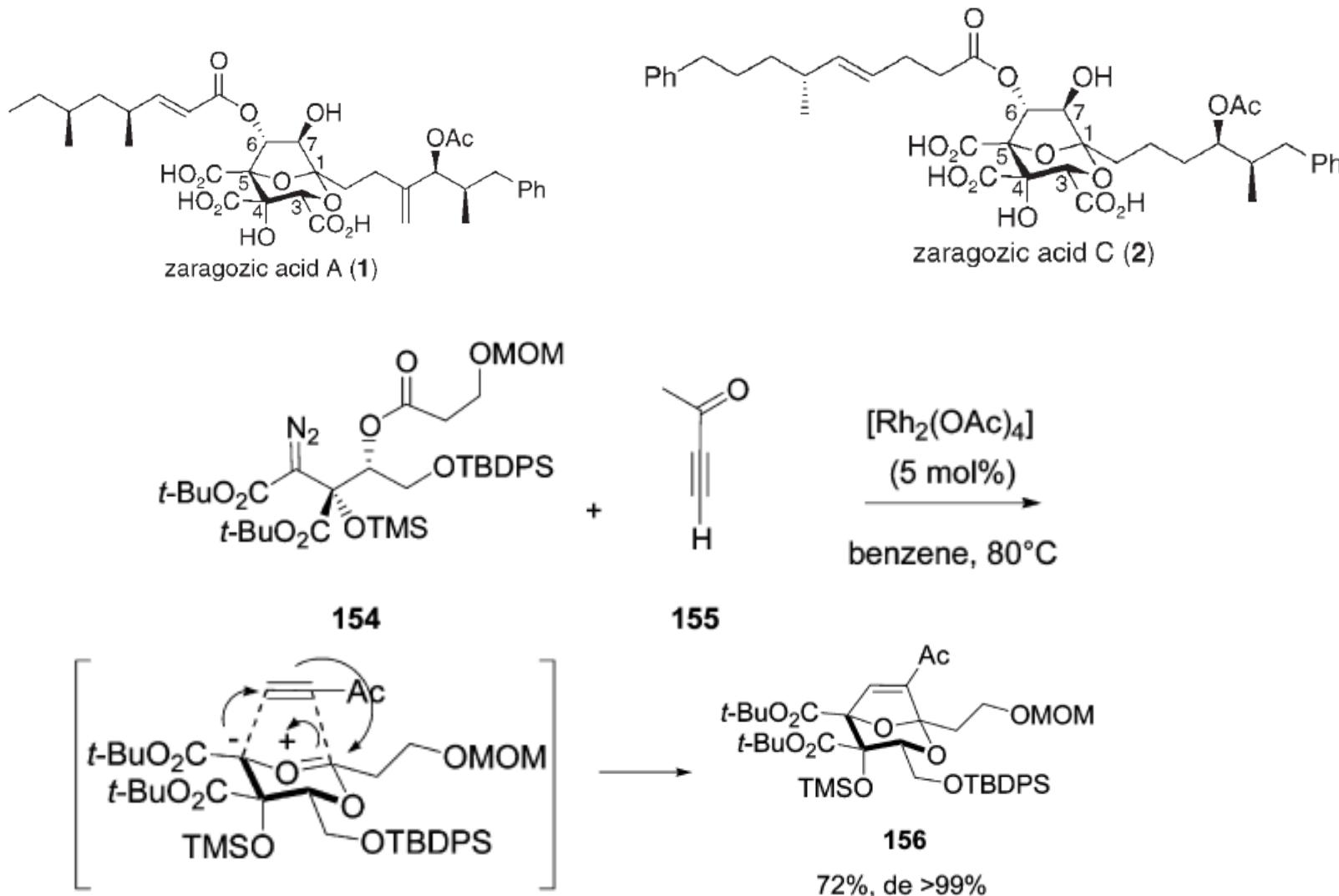


Initiated by a pericyclic primary step

Domino Retro-Diels-Alder Cycloaddition/ Diels-Alder Cycloaddition Reactions

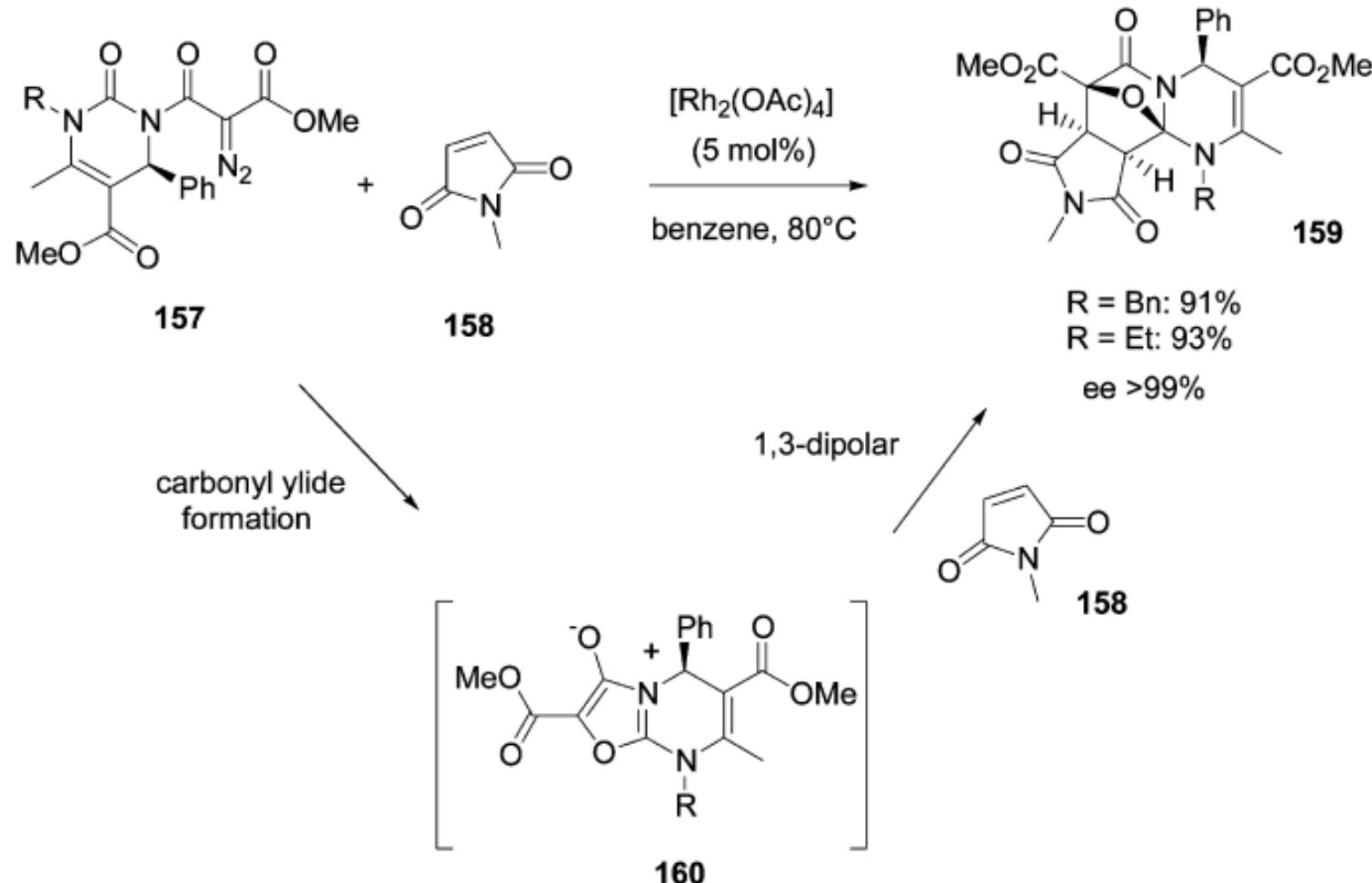


Domino Carbonyl Ylide Formation/1,3-Dipolar Cycloaddition Reaction



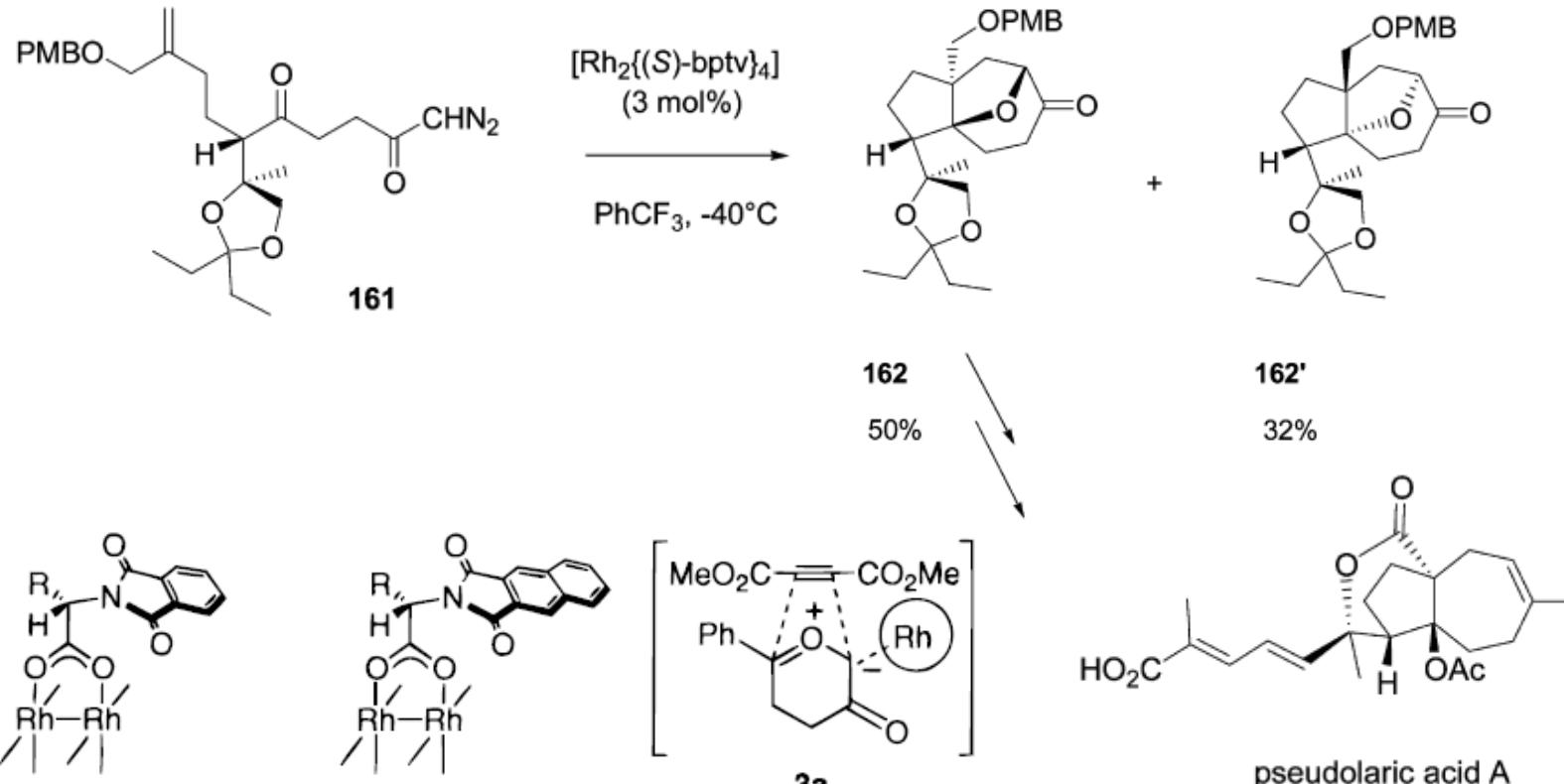
Carbene sequence

Domino Carbonyl Ylide Formation/1,3-Dipolar Cycloaddition Reaction



Carbene sequence

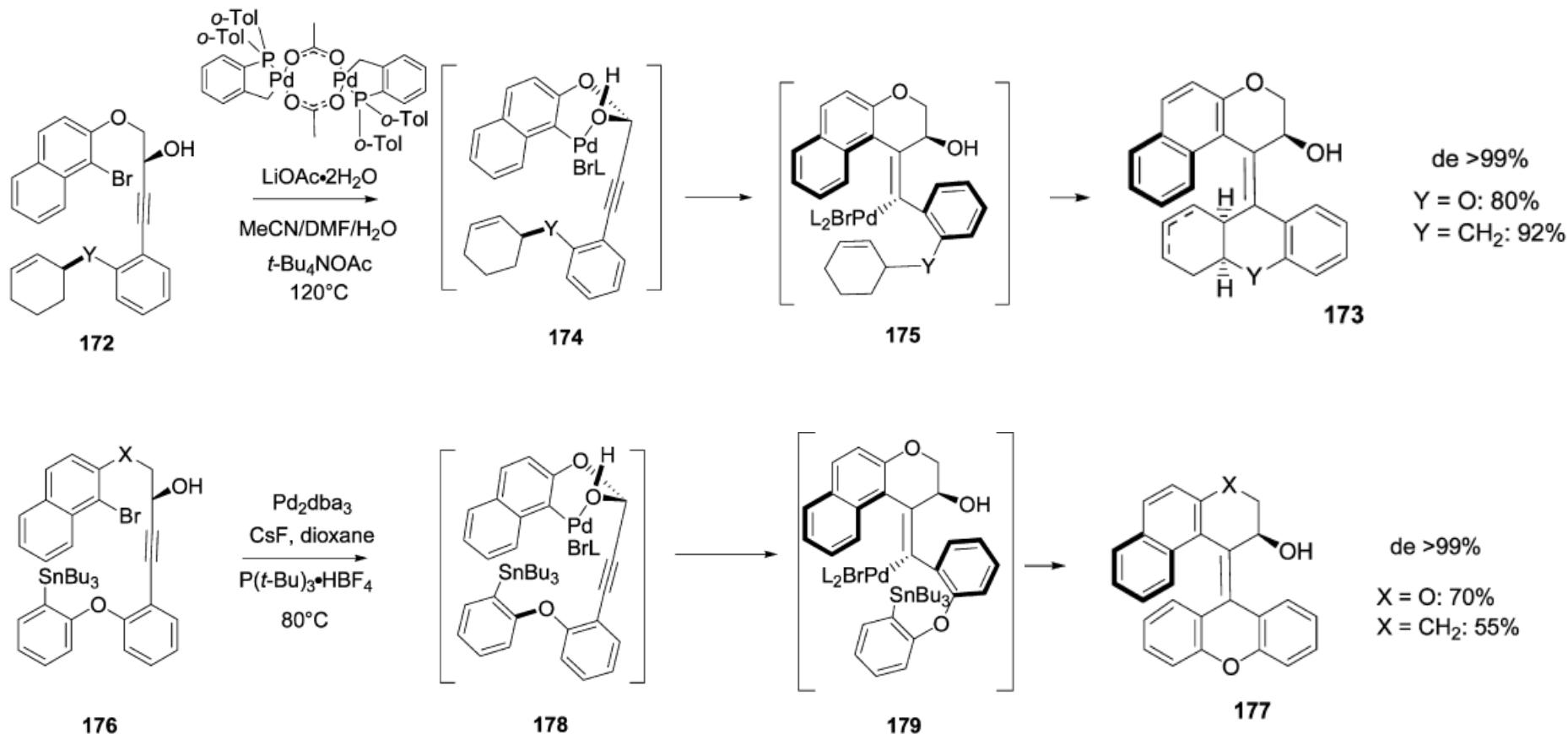
Domino Carbonyl Ylide Formation/1,3-Dipolar Cycloaddition Reaction and Total Synthesis of Pseudolaric Acid A



$\text{R} = \text{Br}: \text{Rh}_2(\text{S-PTPA})_4$ $\text{Rh}_2(\text{S-BPTPA})_4$
 $\text{R} = \text{Me}: \text{Rh}_2(\text{S-PTA})_4$ $\text{Rh}_2(\text{S-BPTA})_4$
 $\text{R} = \text{Pr}^t: \text{Rh}_2(\text{S-PTV})_4$ $\text{Rh}_2(\text{S-BPTV})_4$
 $\text{R} = \text{Bu}^t: \text{Rh}_2(\text{S-PTTL})_4$ $\text{Rh}_2(\text{S-BPTTL})_4$

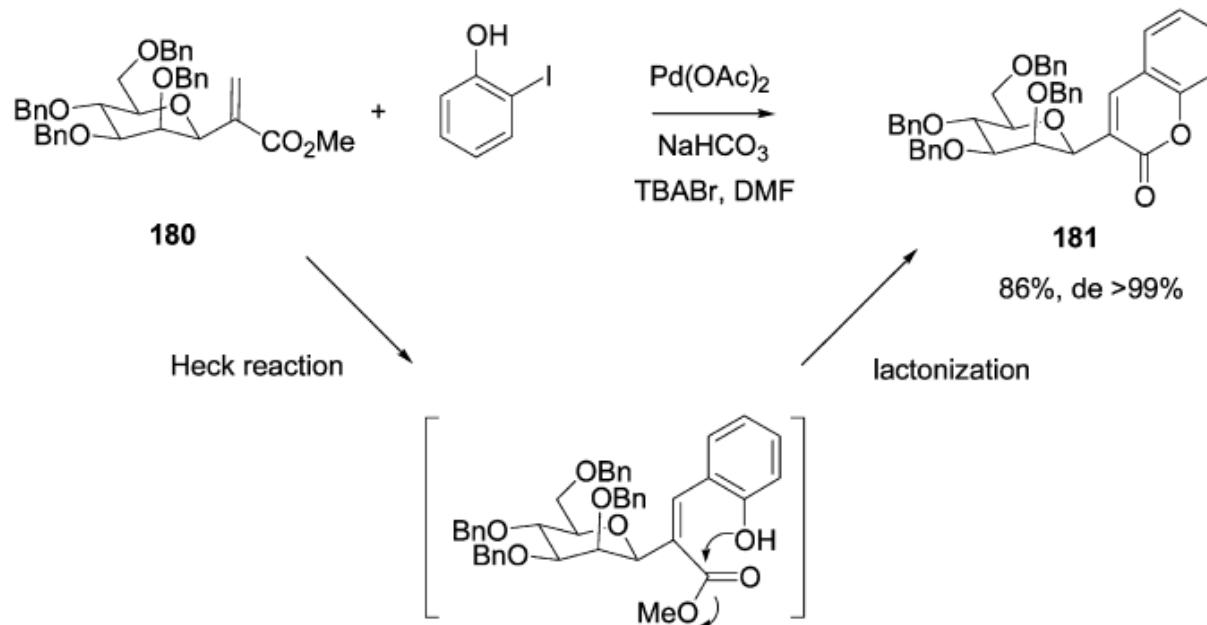
Palladium catalyzed domino reaction

Domino Carbopalladation/Heck Reaction and Domino Carbopalladation/Stille Reaction

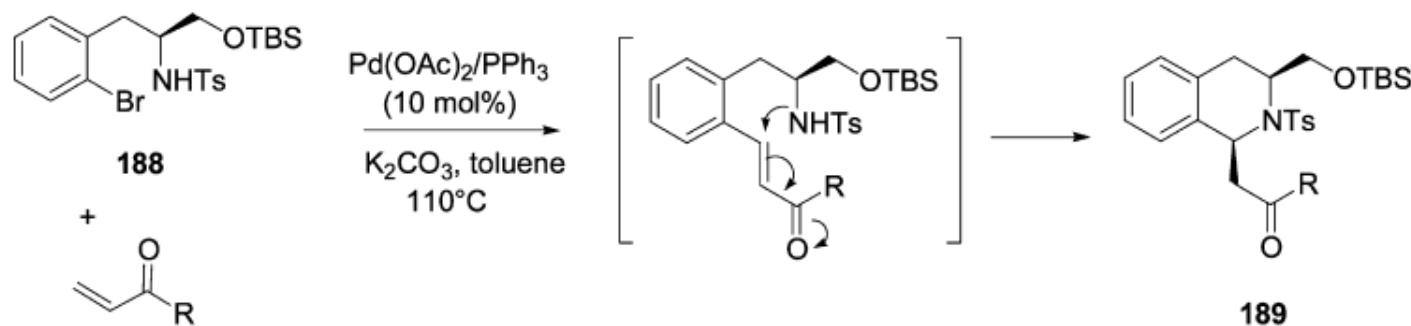


Palladium catalyzed domino reaction

Domino Heck/Lactonization Reaction



Domino Heck/Aza-Michael Reaction

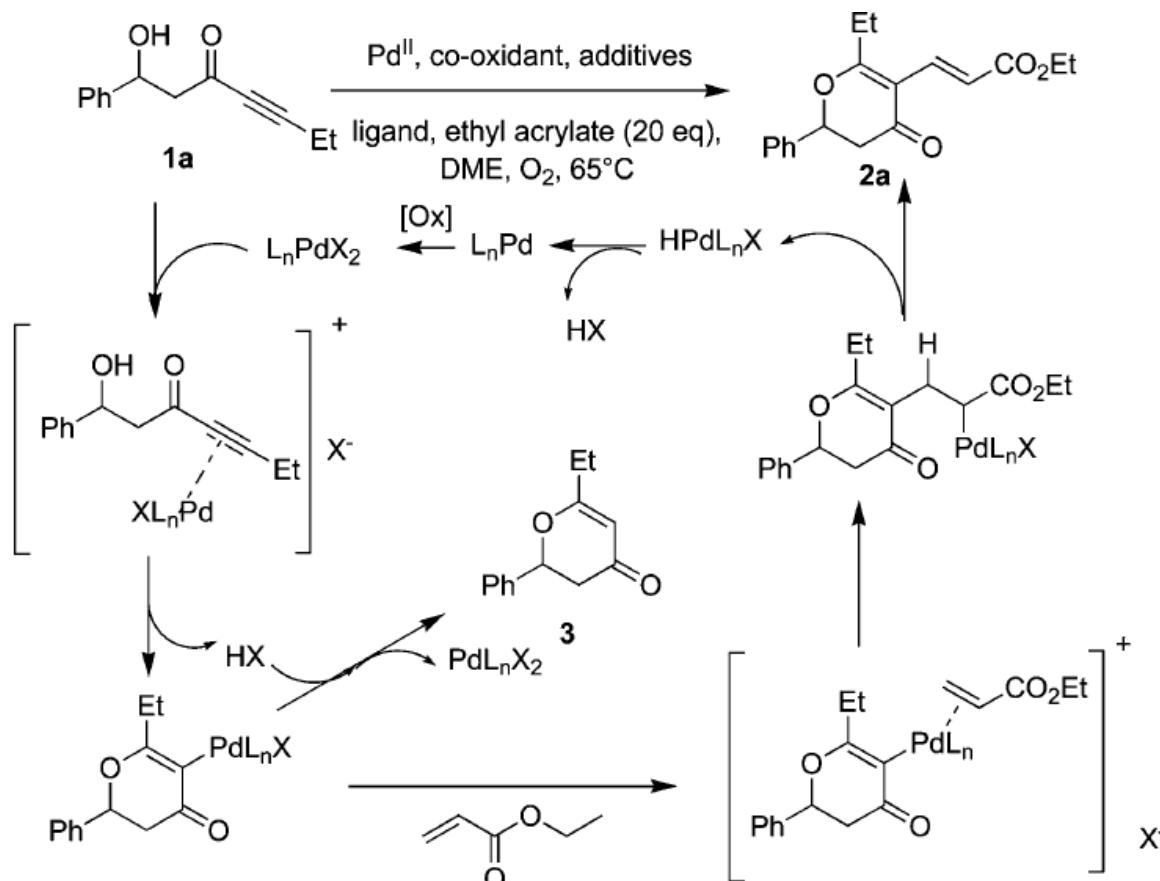
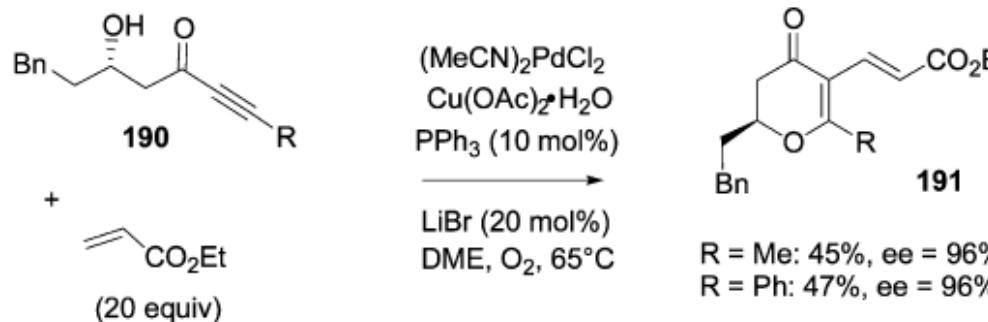


Roy, R. J. Org. Chem. 2009, 74, 8480

Pfeffer, F. M. Tetrahedron Lett. 2012, 53, 1468

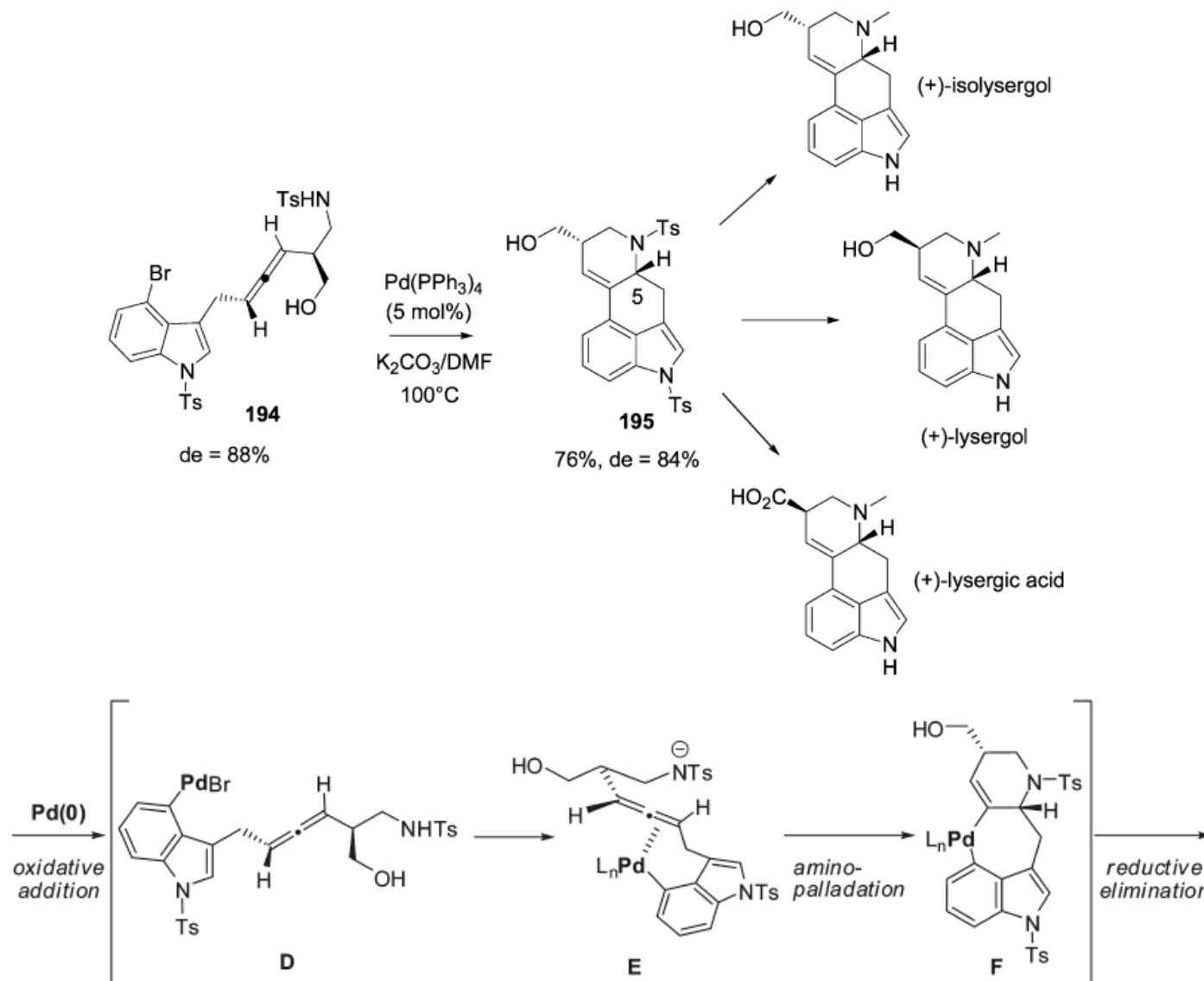
Palladium catalyzed domino reaction

Domino Wacker/Heck Reaction



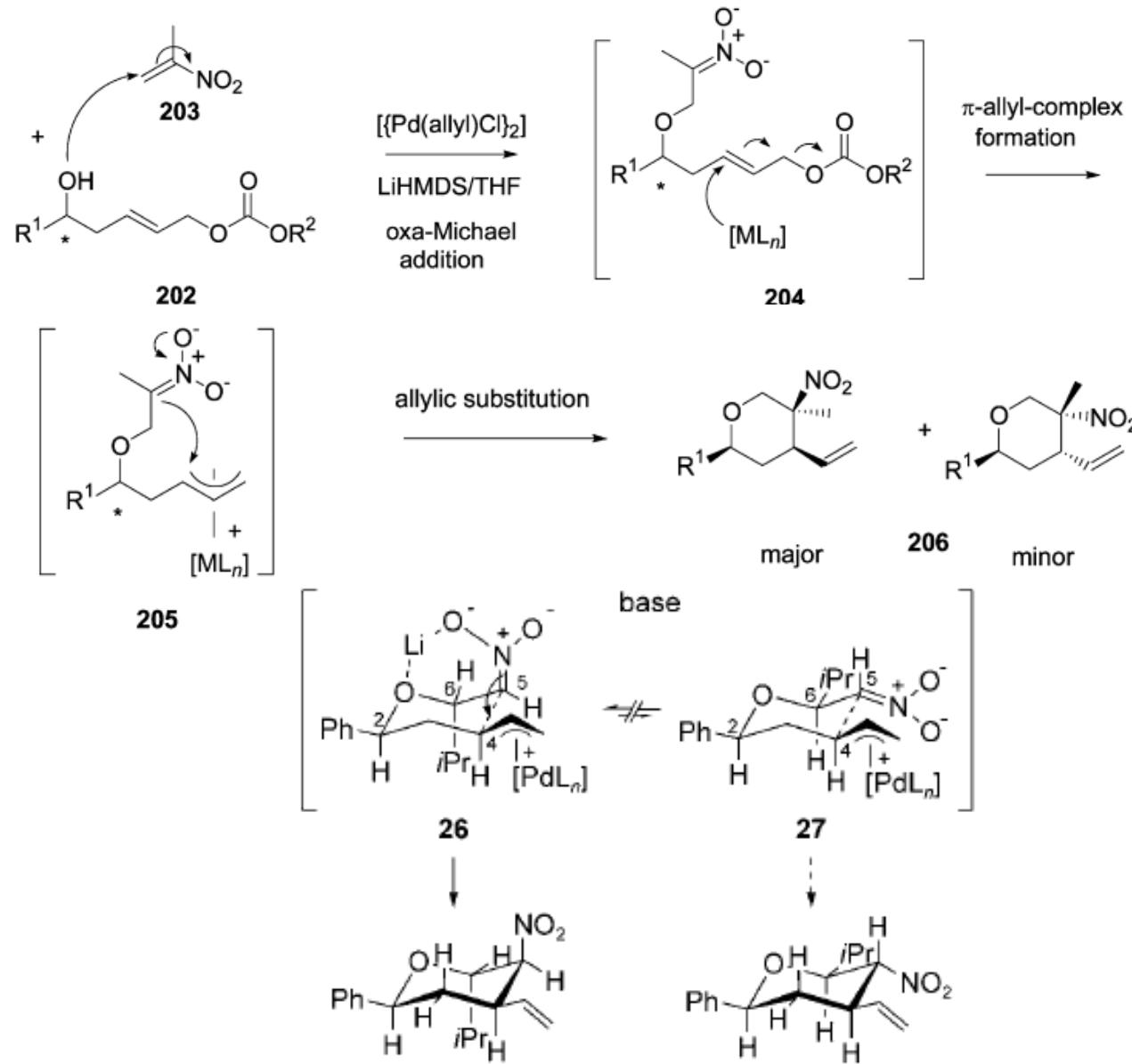
Palladium catalyzed domino reaction

Syntheses of (+)-Lysergic Acid, (+)-Lysergol, and (+)-Isolysergol through Domino Cyclization Reaction



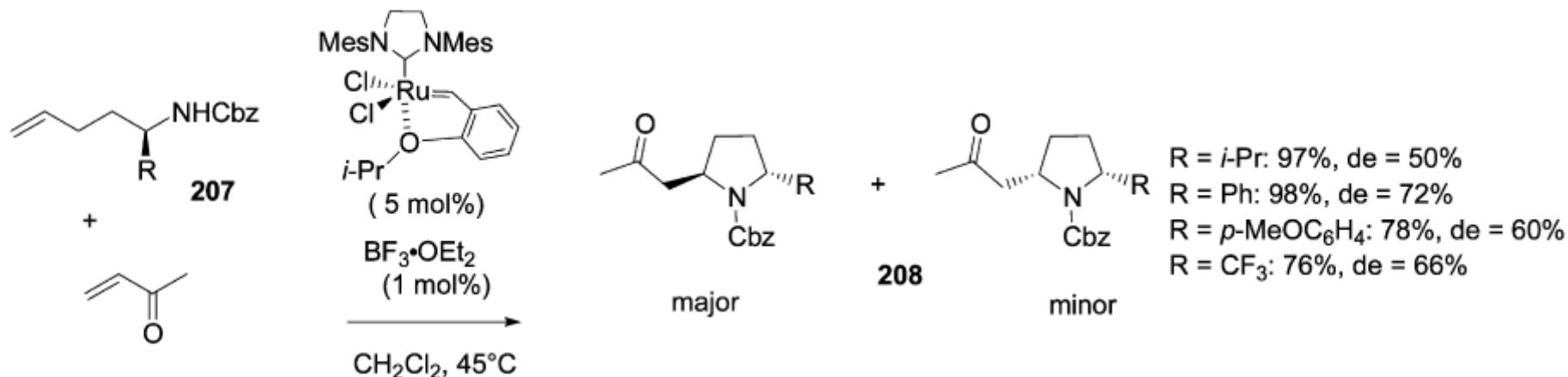
Palladium catalyzed domino reaction

Domino Oxa-Michael/Tsuji-Trost Reaction

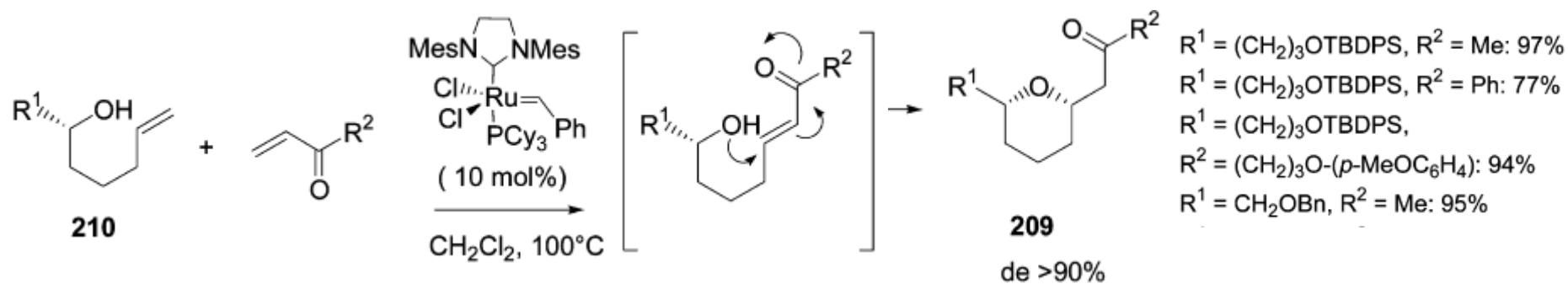


Ruthenium catalyzed domino reaction

Domino Olefin Cross-Metathesis/Intramolecular Aza-Michael Reaction



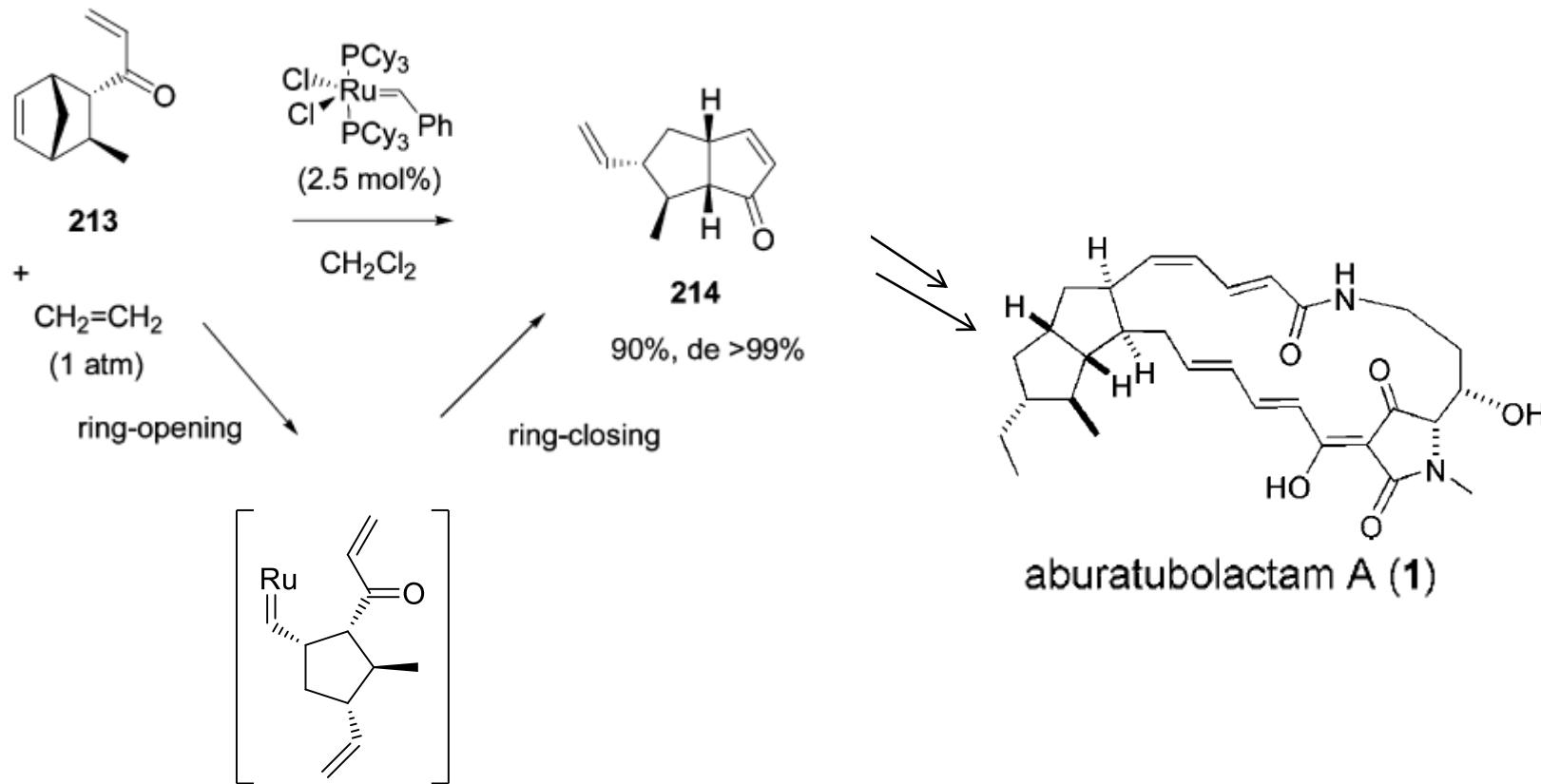
Domino Olefin Cross-Metathesis/Intramolecular Oxa-Michael Reaction



Pozo, C. *J. Am. Chem. Soc.* **2007**, *129*, 6700

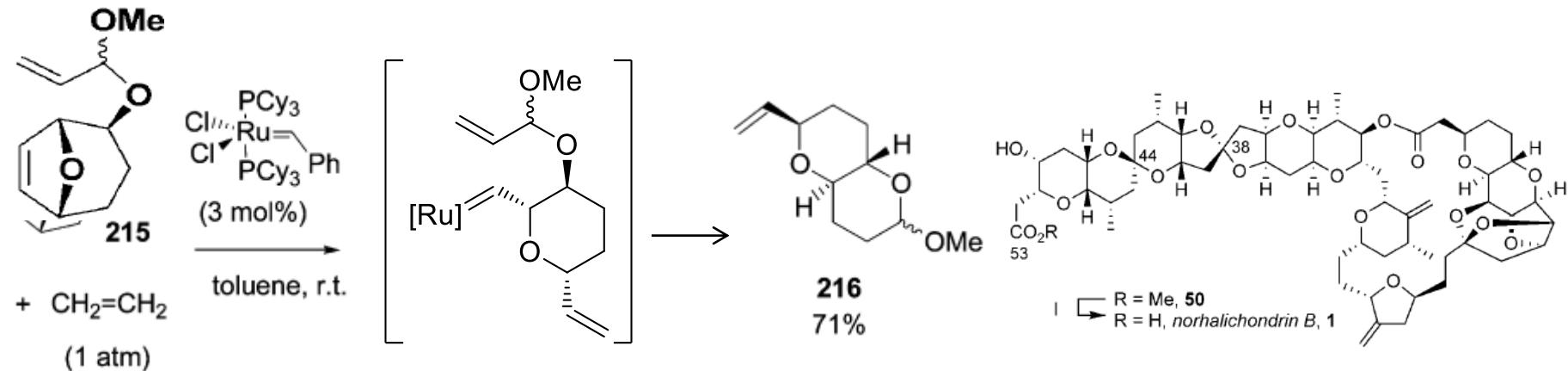
Sasaki, M. *Org. Lett.* **2010**, *12*, 1636

Domino Ring-Opening/Ring-Closing Metathesis Reactions

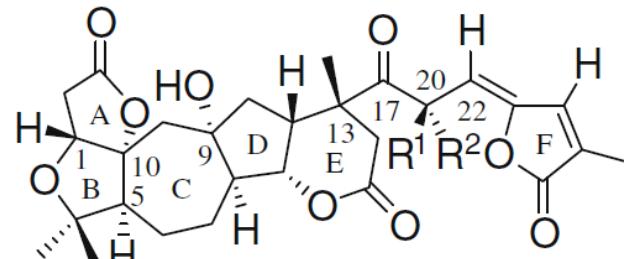
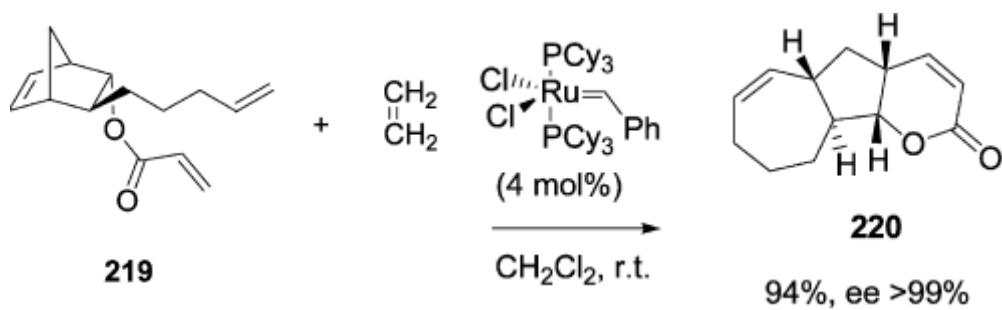


Ruthenium catalyzed domino reaction

Domino Ring-Opening/Ring-Closing Metathesis Reactions



Phillips, A. J. *Angew. Chem., Int. Ed.* **2009**, *48*, 2346

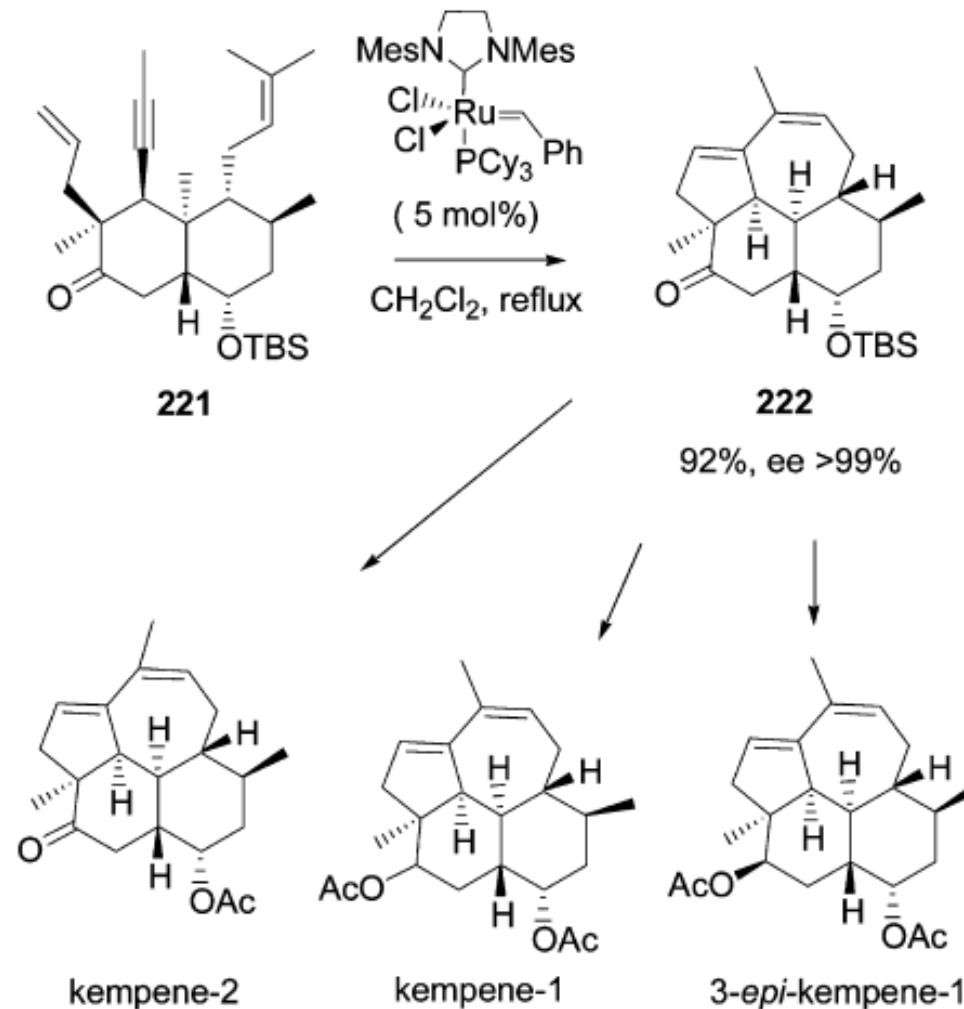


schintrilactones

Ghosh, S. *Tetrahedron Lett.* **2010**, *51*, 2754

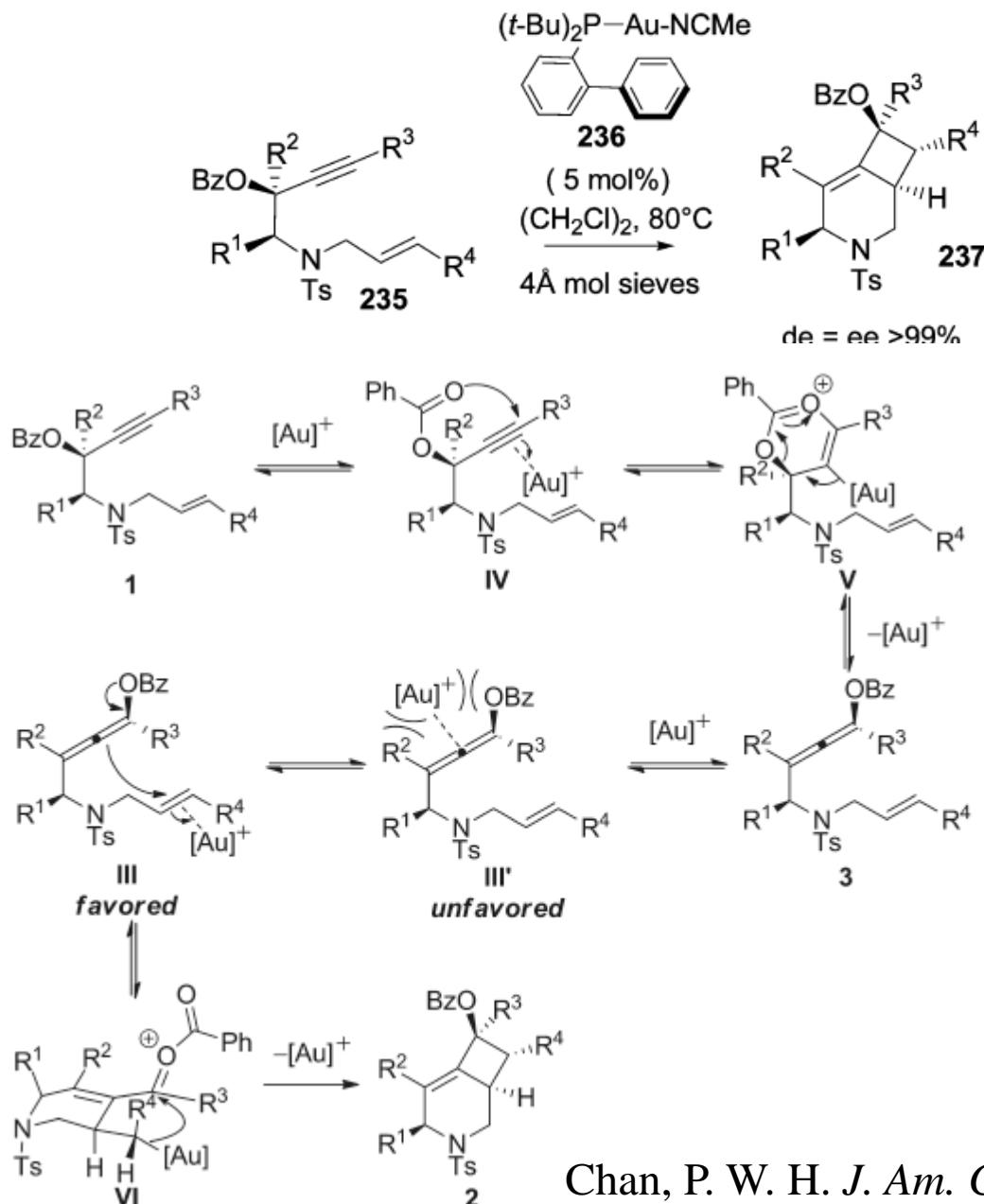
Ruthenium catalyzed domino reaction

Synthesis of Kempene-2, Kempene-1, and 3-*epi*-Kempene-1 through Domino Enyne Metathesis Reaction



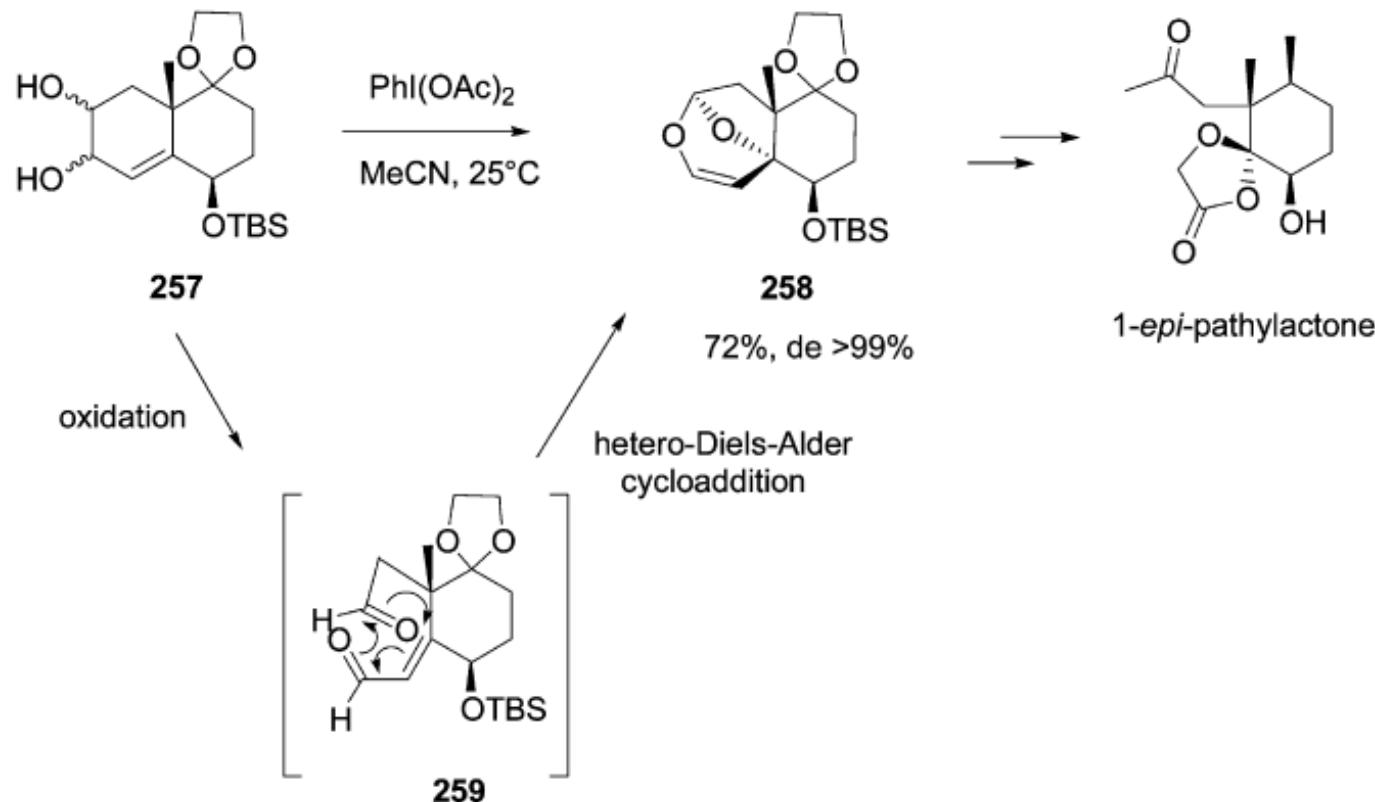
Miscellaneous domino reactions

Domino 1,3-Migration/[2 + 2]-Cycloaddition Reaction



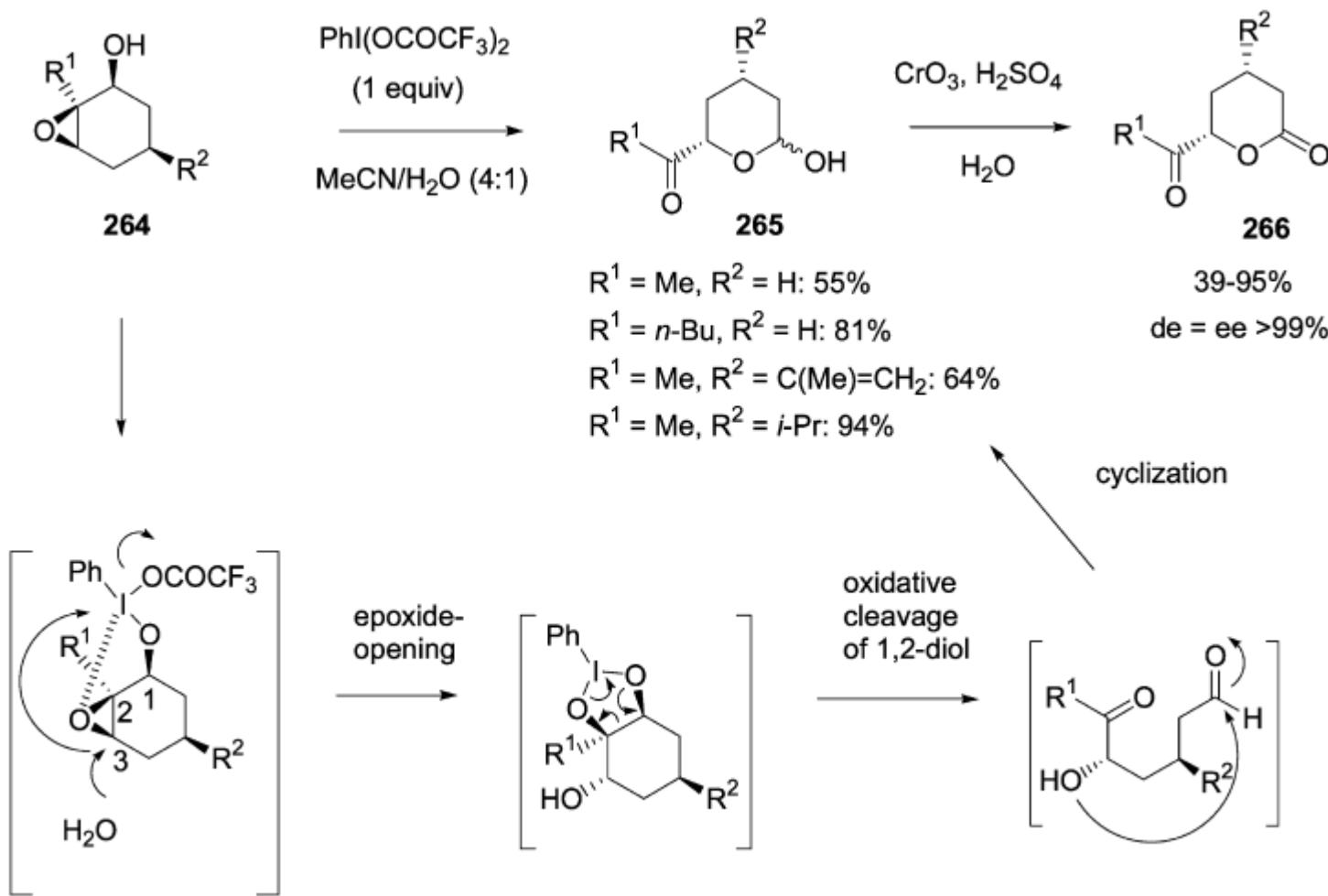
Miscellaneous domino reactions

Domino Oxidation/Intramolecular Hetero-Diels-Alder Cycloaddition Reaction

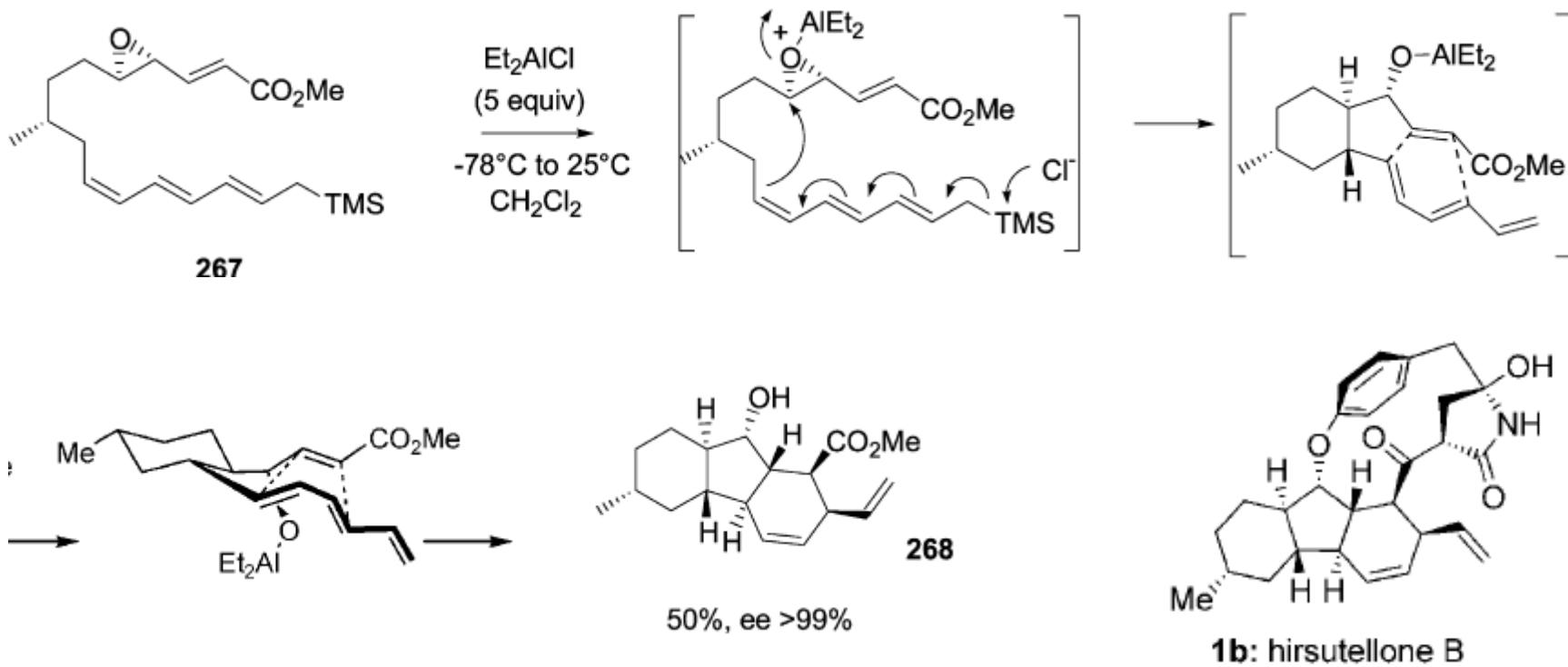


Miscellaneous domino reactions

Domino Epoxide-Opening/Oxidative Cleavage/Cyclization Reaction

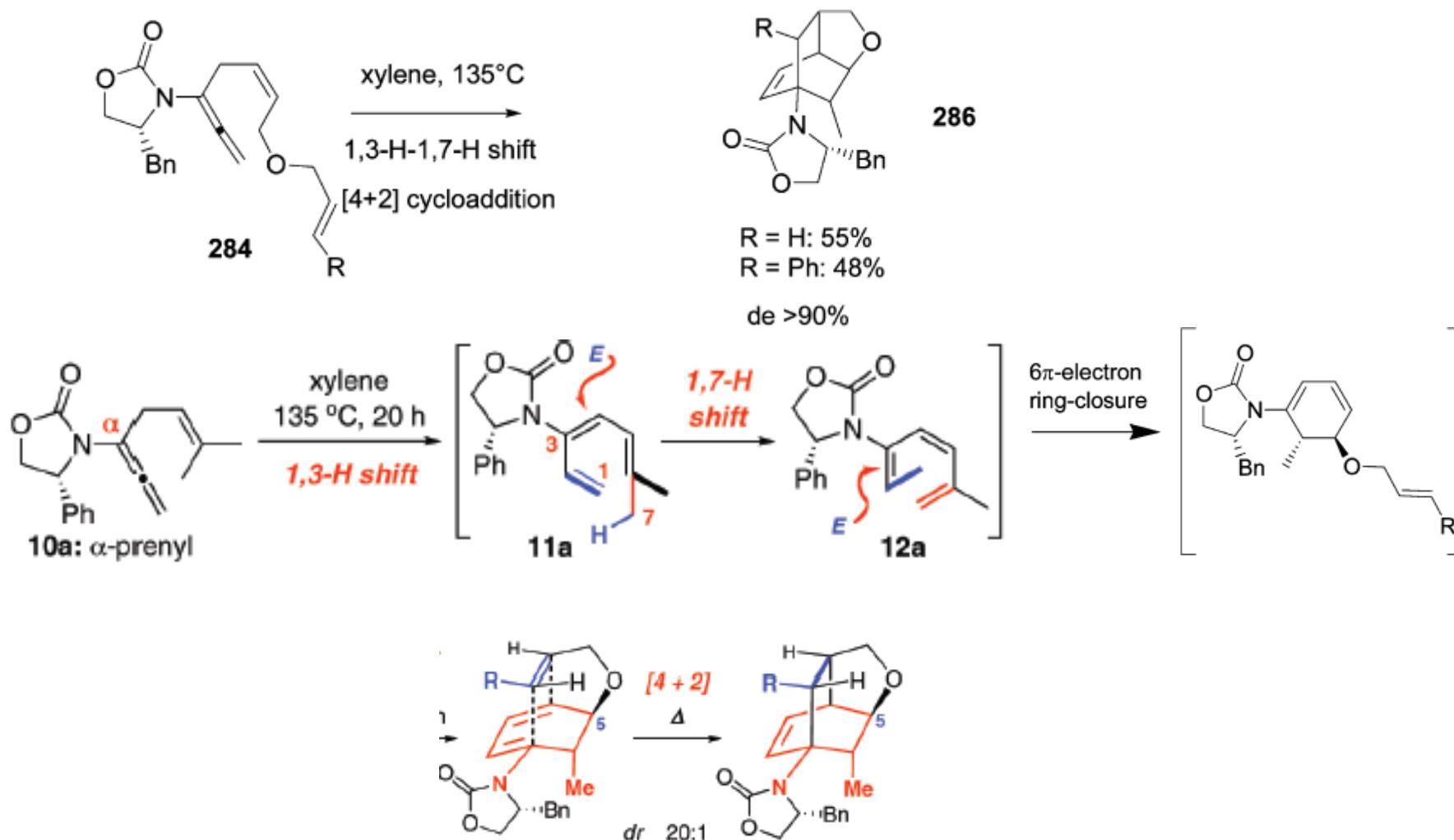


Synthesis of Hirsutellone B through Al-Catalyzed Domino Epoxide-Opening/Diels-Alder Reaction



Miscellaneous domino reactions

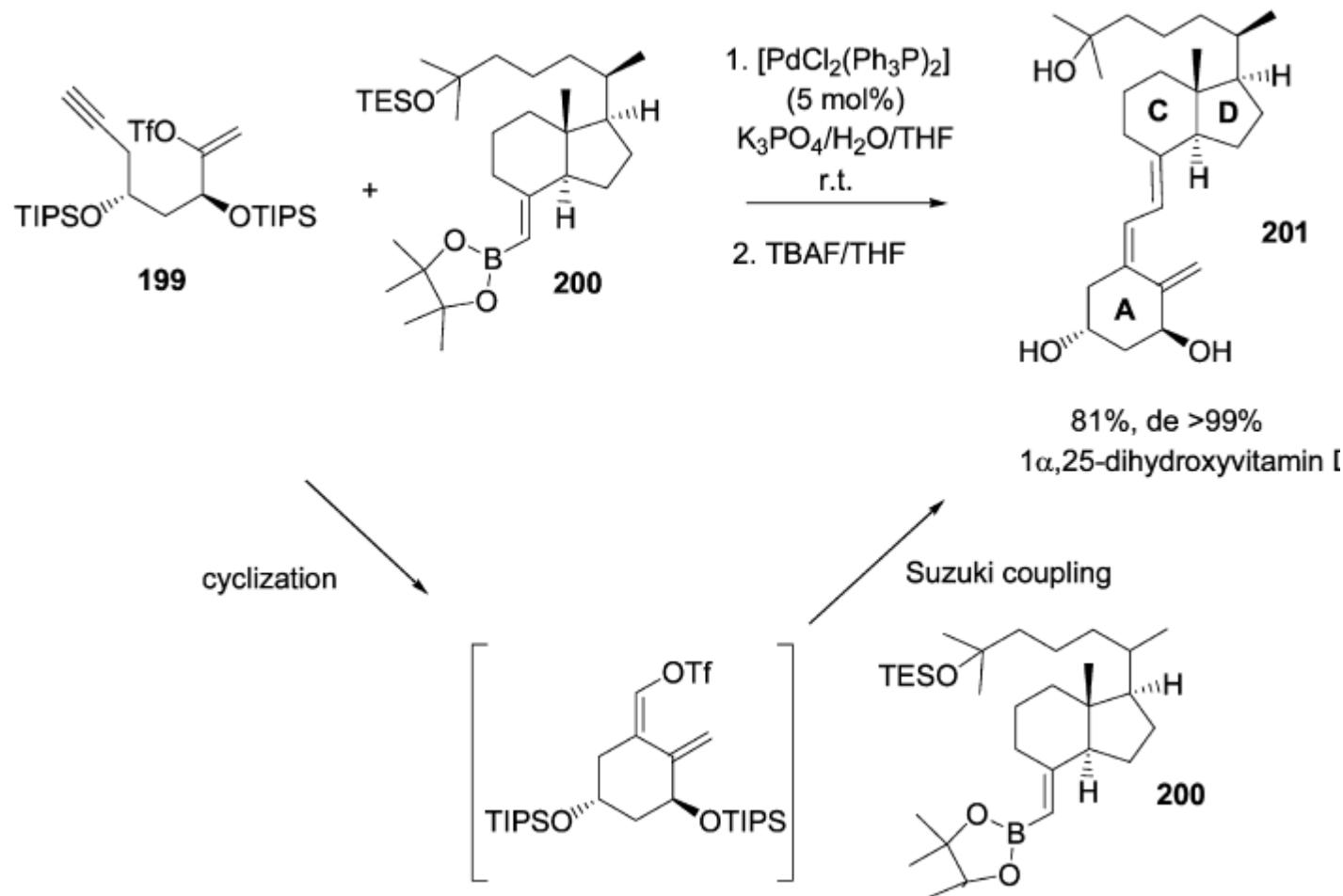
Domino 1,3-H-1,7-H Shift/6 π Electron Pericyclic Ring-Closure/[4 + 2]-Cycloaddition Reaction





Palladium catalyzed domino reaction

Synthesis of $1\alpha,25$ -Dihydroxyvitamin D₃ through Domino Cyclization/Suzuki Coupling Reaction



Domino bromination/Cyclization Reactions

