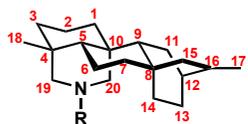
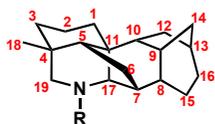


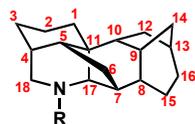
## Skeletons of diterpenoid alkaloids



C20-diterpenoid alkaloids



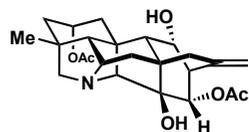
C19-diterpenoid alkaloids



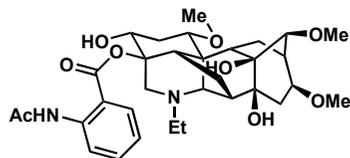
C18-diterpenoid alkaloids

Isolation: *Aconitum*, *Delphinium*, *Consolida*, *Spiraea*...  
over 1200 species

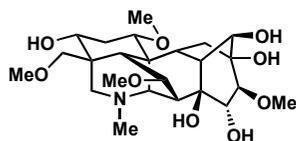
## Representative members of the diterpenoid alkaloids



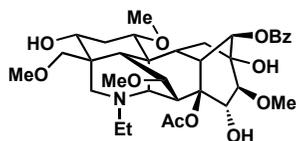
guan-fu base A [C<sub>20</sub>]  
antiarrhythmic  
Na<sub>v</sub> blocker



lappaconitine [C<sub>18</sub>]  
analgesic  
Na<sub>v</sub> blocker

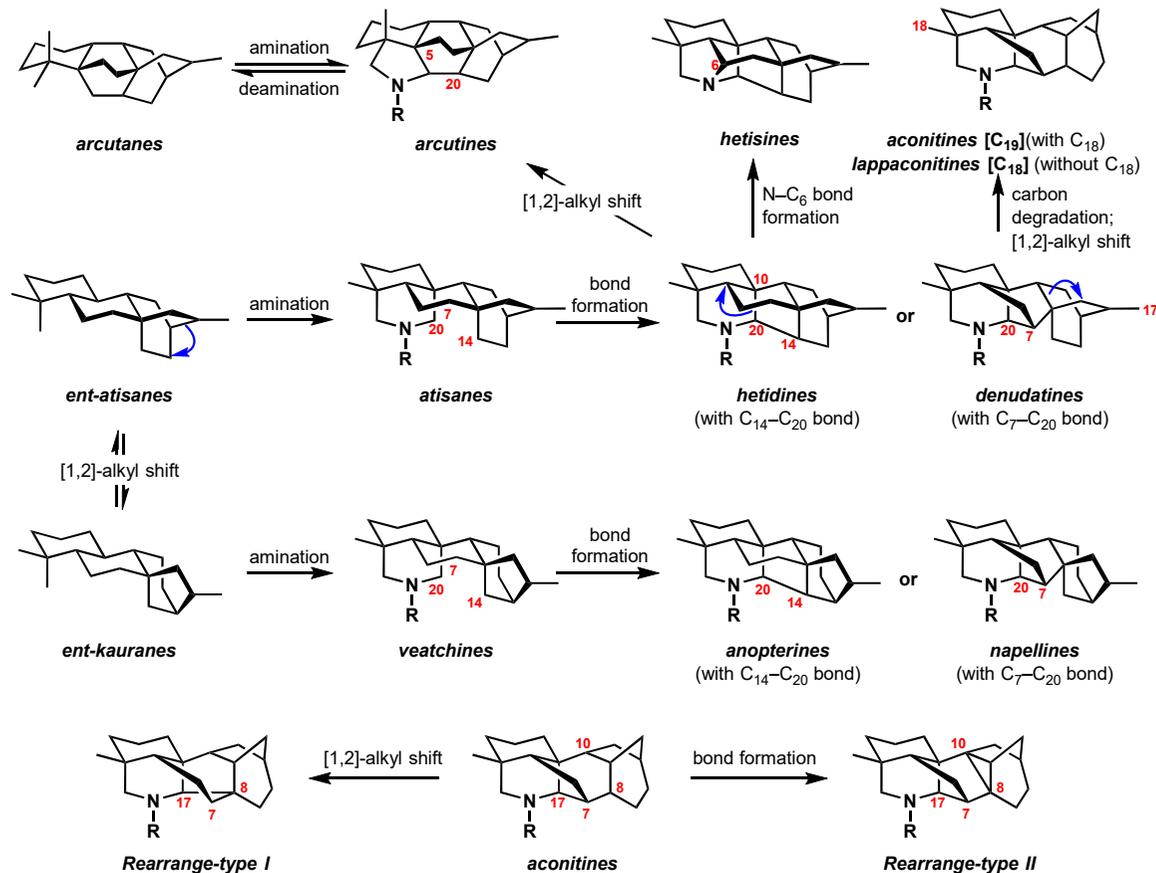


mesaconine [C<sub>19</sub>]  
cardiotonic  
Na<sub>v</sub> activator



aconitine [C<sub>19</sub>]  
highly toxic  
Na<sub>v</sub> activator

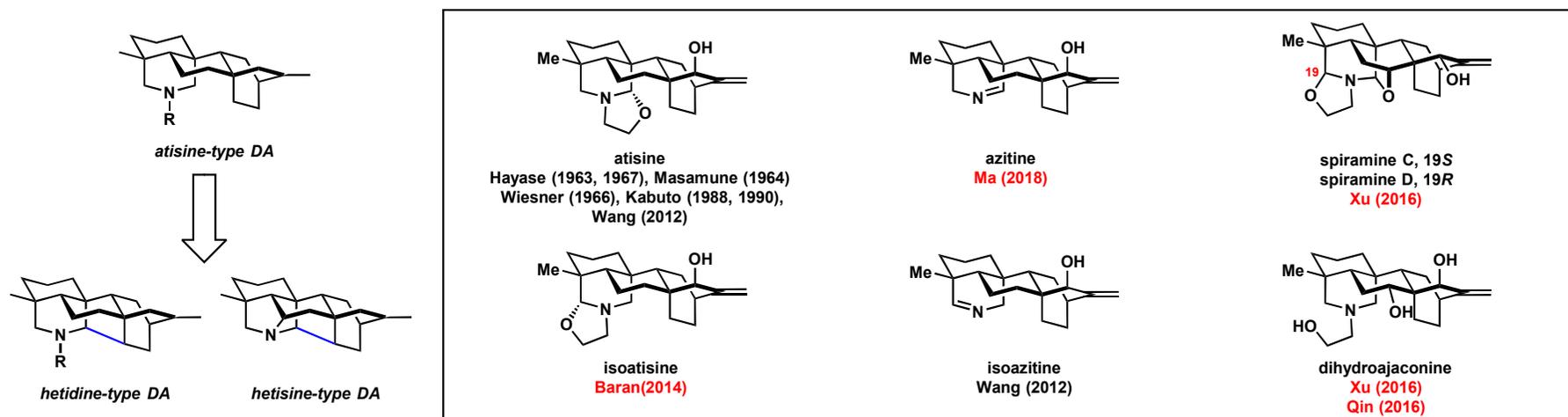
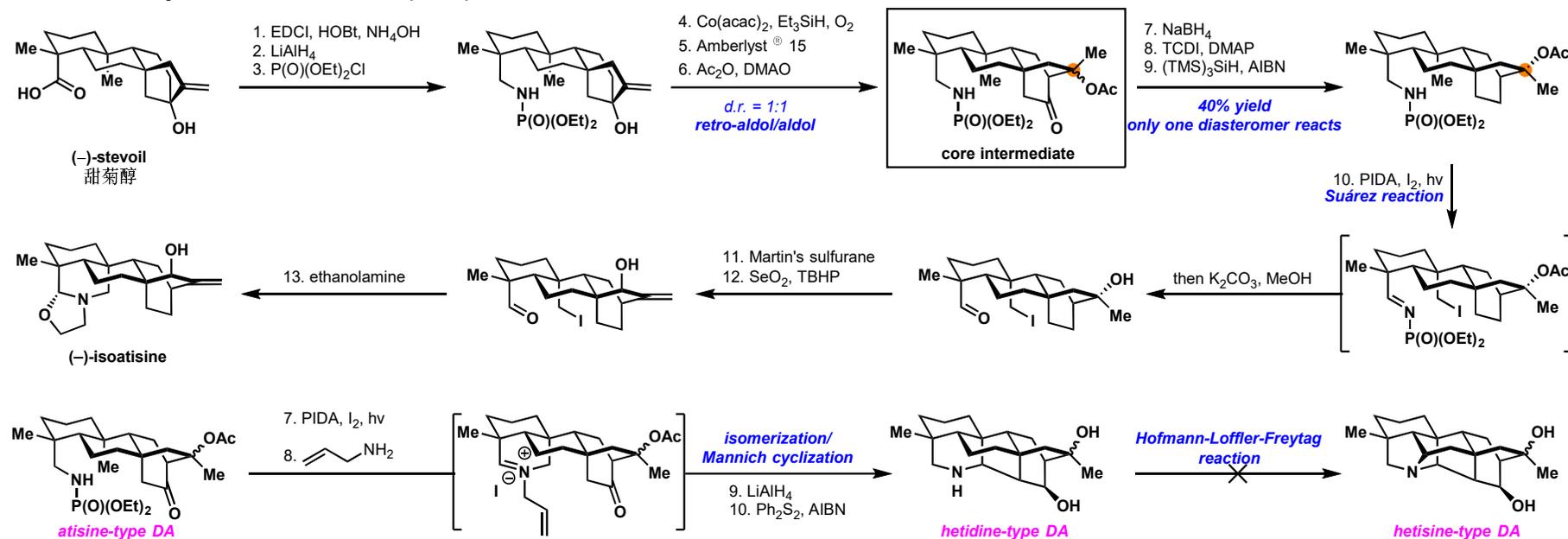
## Biosynthetic Relationships of Diterpenoid Alkaloids



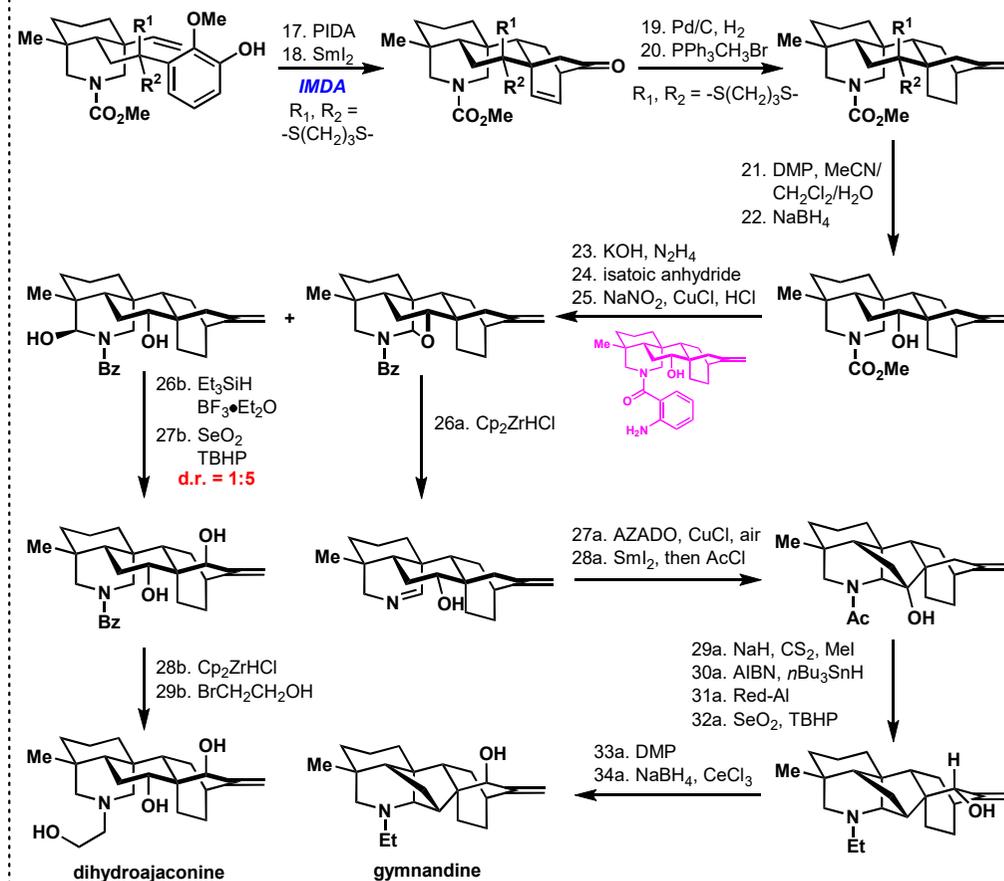
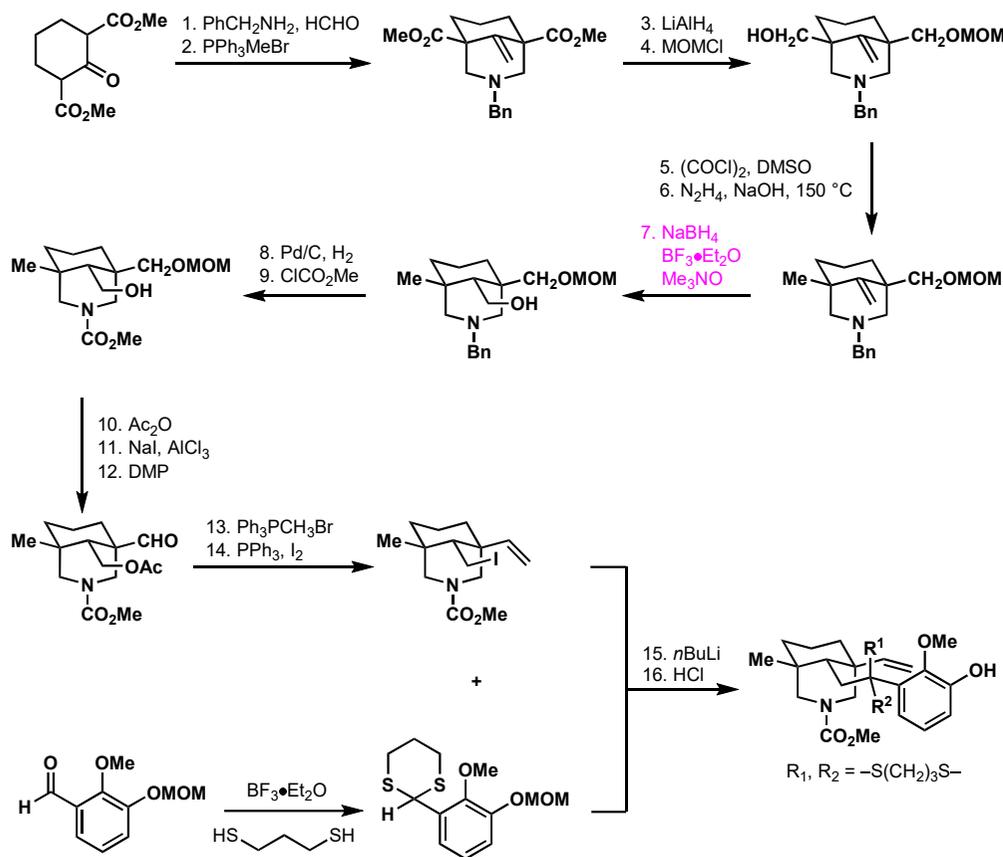
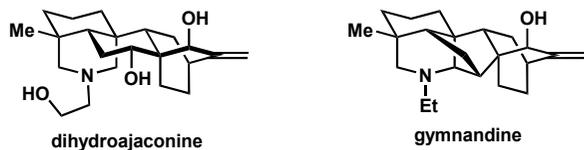
## These are reviews you MUST READ:

- F. Wang *et al.* *Nat. Prod. Rep.* **2010**, 27, 528
- B. Liu *et al.* *Synthesis* **2015**, 47, 2691
- Y. Qin *et al.* *Nat. Prod. Rep.* **2017**, 34, 1044
- M. Lautens *et al.* *Synthesis* **2019**, 51, 3915
- R. Sarpong, R. W. Hoffmann *et al.* *Angew. Chem. Int. Ed.* **2020**, 59, 10722
- R. Sarpong *et al.* *Chem. Sci.* **2020**, 11, 7538
- Y. Qin *et al.* *Acc. Chem. Res.* **2021**, 54, 22

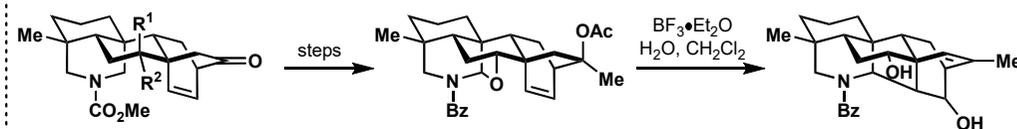
## Total Synthesis of the Atisine-type Alkaloids

Baran's Synthesis of Isoatisine (2014) P. S. Baran *et al.* *J. Am. Chem. Soc.* 2014, 136, 12592

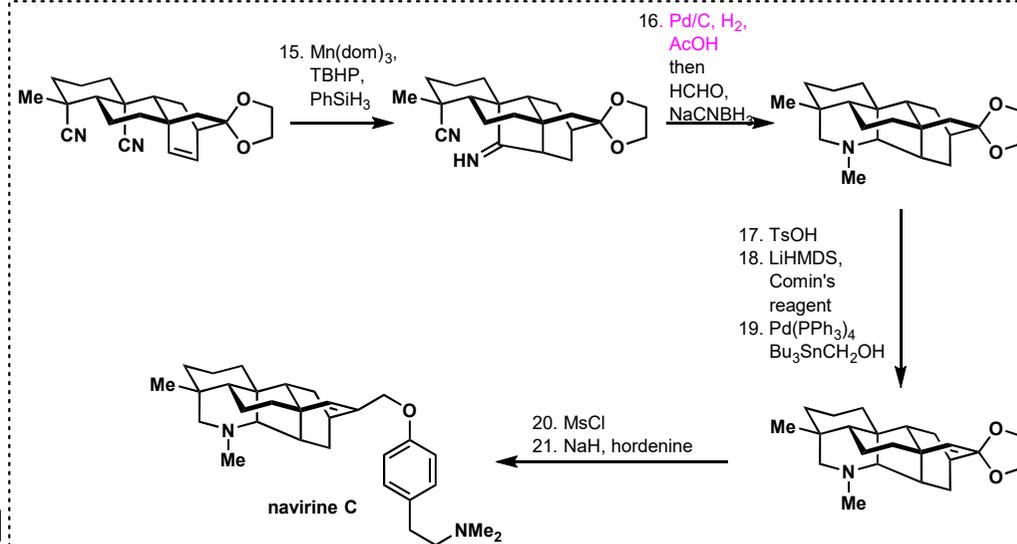
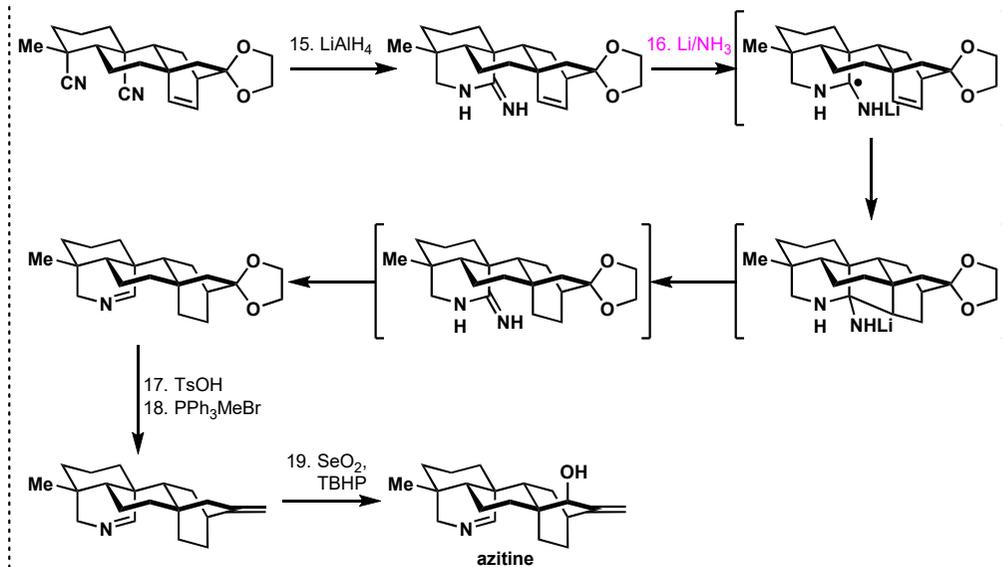
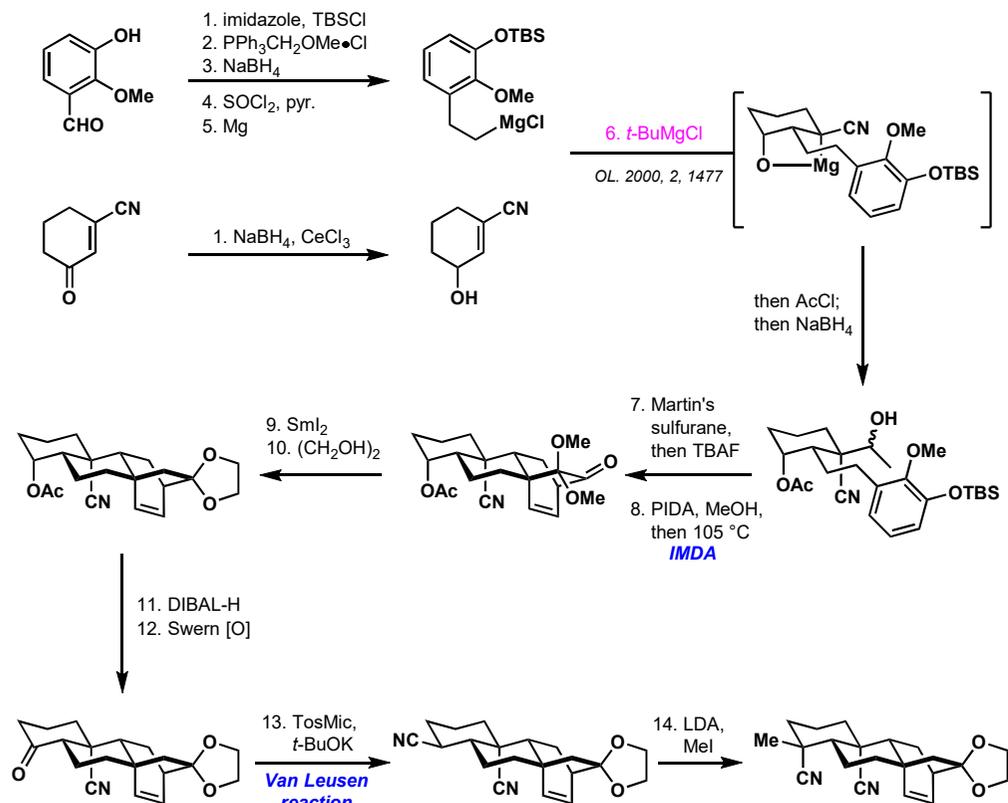
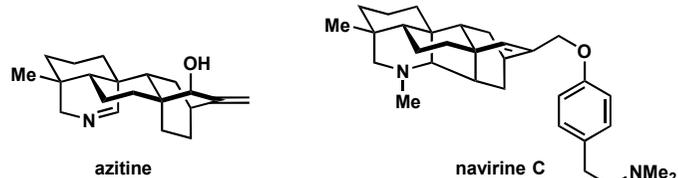
## Qin's synthesis of dihydroajaconine and gymnandine

Y. Qin *et al.* *Angew. Chem. Int. Ed.* **2016**, *55*, 392

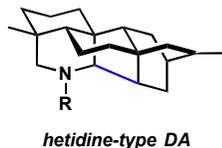
## Qin's hetidine skeleton synthesis



## Ma's synthesis of atizine and navirine C

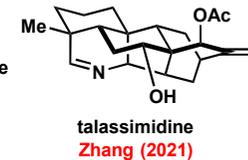
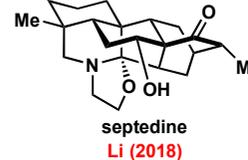
D. Ma et al. *Angew. Chem. Int. Ed.* 2018, 57, 6676

## Total Synthesis of the Hetidine-type Alkaloids



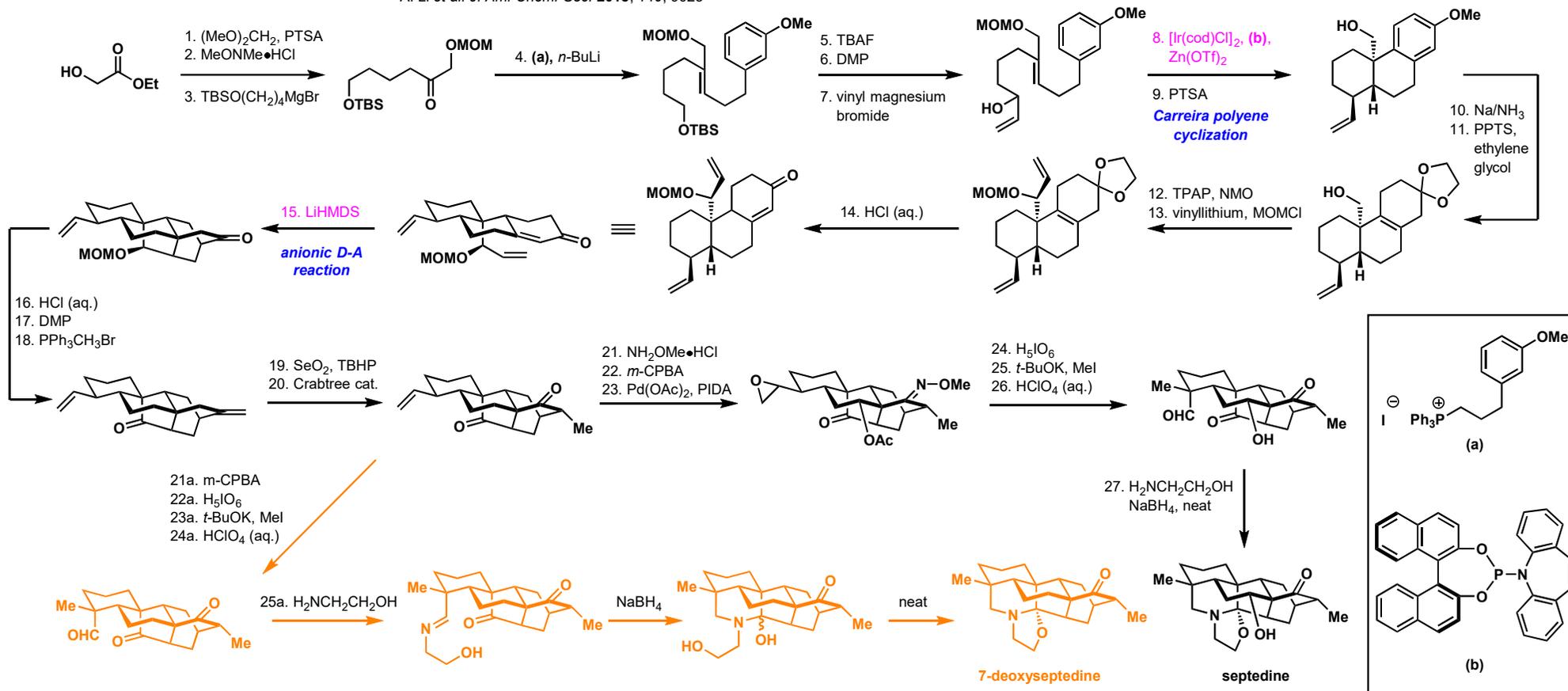
## skeleton synthesis

R. Sarpong *et al.* *Angew. Chem., Int. Ed.* **2013**, 52, 4854  
 P. S. Baran *et al.* *J. Am. Chem. Soc.* **2014**, 136, 12592  
 Y. Qin *et al.* *Angew. Chem. Int. Ed.* **2016**, 55, 392  
 D. Ma *et al.* *Angew. Chem. Int. Ed.* **2018**, 57, 6676  
 M. Zhang *et al.* *Angew. Chem. Int. Ed.* **2018**, 57, 937

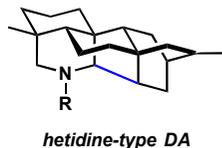


## Li's Synthesis of septedine

A. Li *et al.* *J. Am. Chem. Soc.* **2018**, 140, 9025

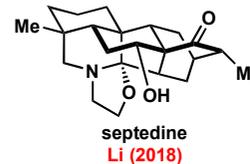


## Total Synthesis of the Hetidine-type Alkaloids



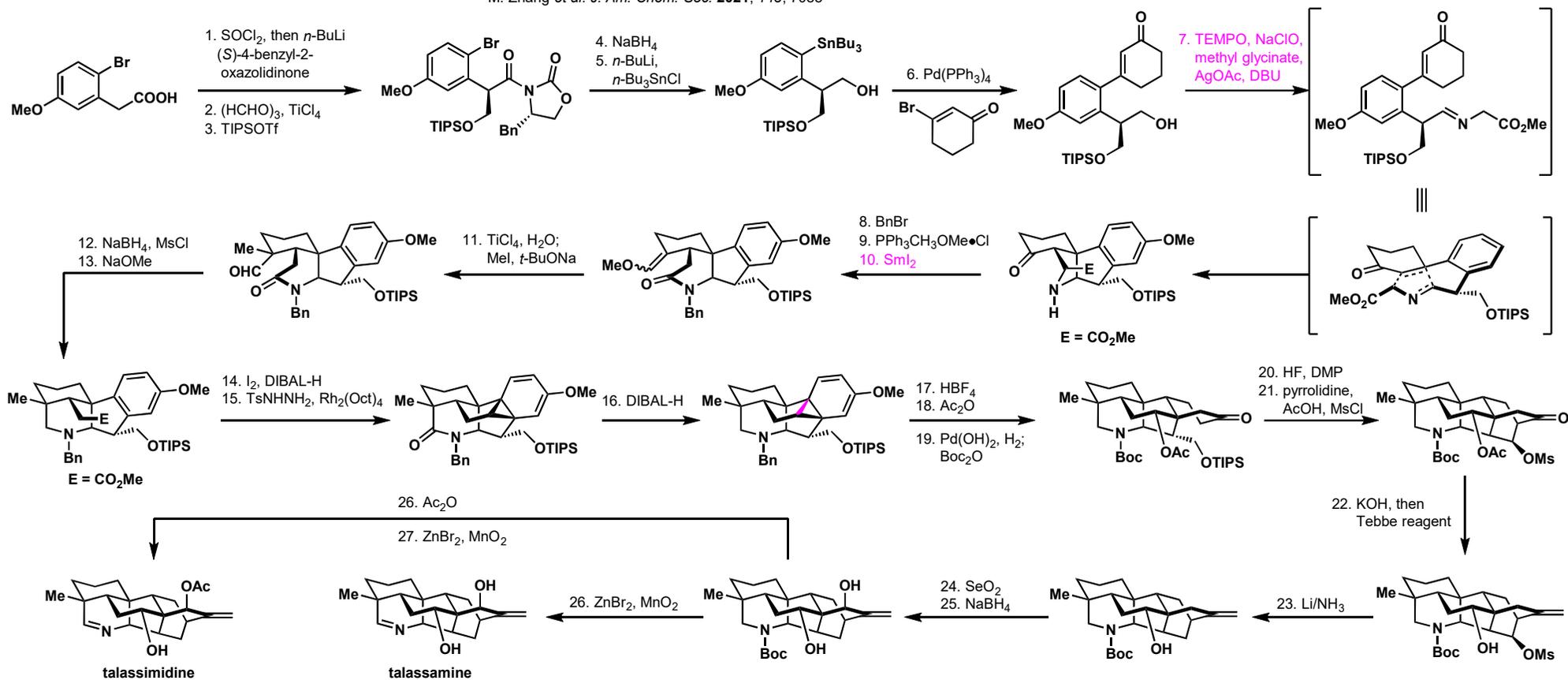
## skeleton synthesis

R. Sarpong *et al.* *Angew. Chem., Int. Ed.* **2013**, 52, 4854  
 P. S. Baran *et al.* *J. Am. Chem. Soc.* **2014**, 136, 12592  
 Y. Qin *et al.* *Angew. Chem. Int. Ed.* **2016**, 55, 392  
 D. Ma *et al.* *Angew. Chem. Int. Ed.* **2018**, 57, 6676  
 M. Zhang *et al.* *Angew. Chem. Int. Ed.* **2018**, 57, 937

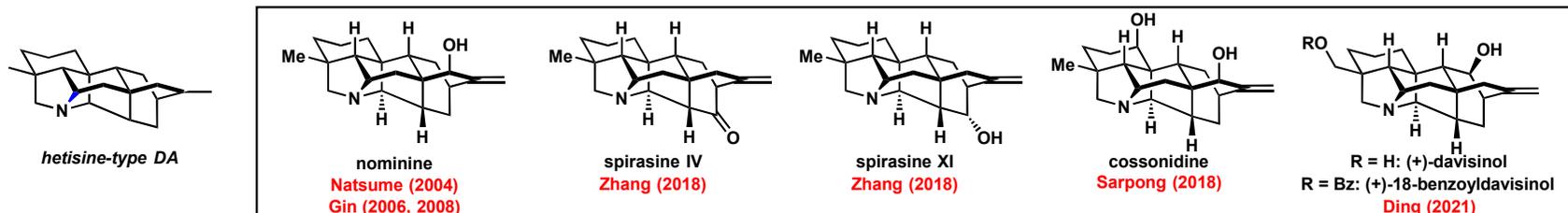


## Zhang's synthesis of talassimidine and talassamine

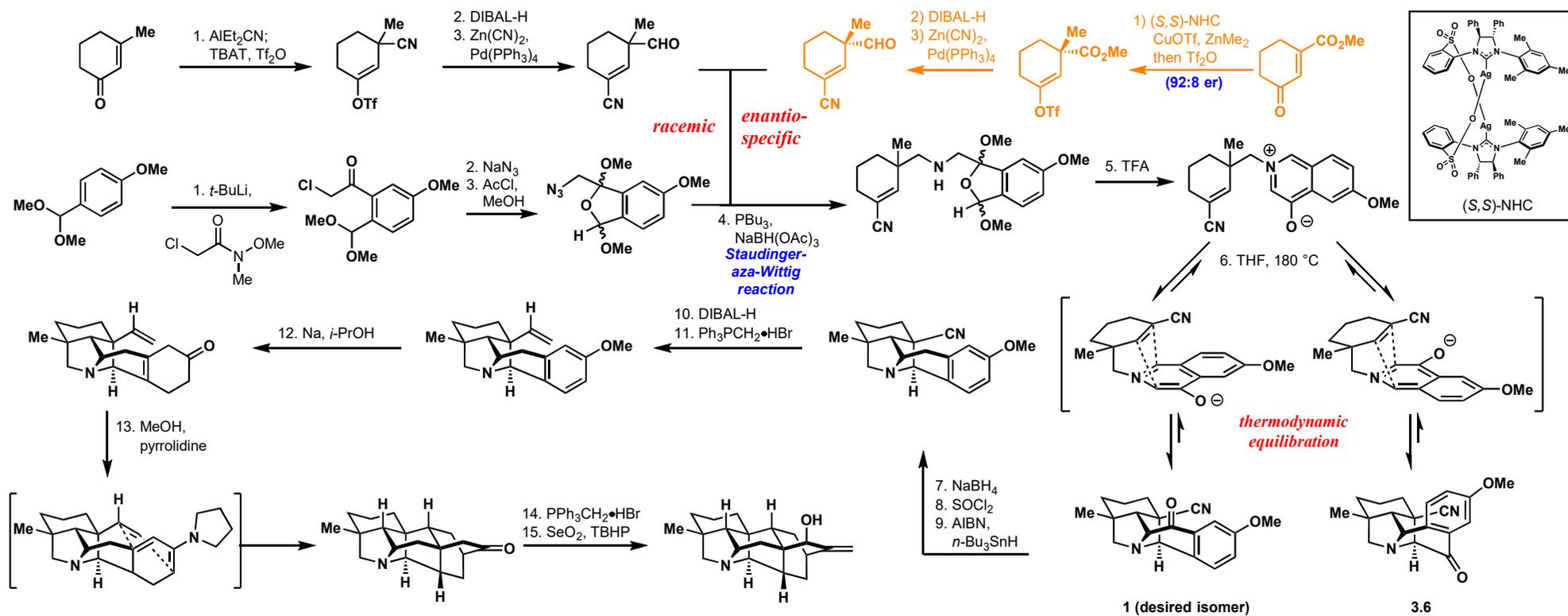
M. Zhang *et al.* *J. Am. Chem. Soc.* **2021**, 143, 7088



## Total Synthesis of the Hetisine-type Alkaloids

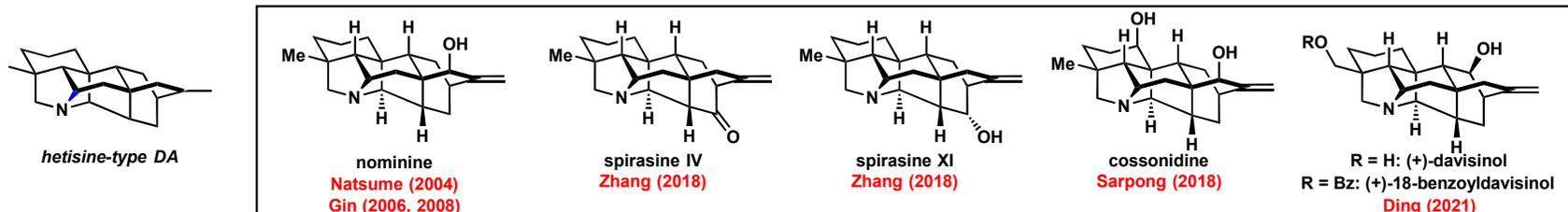


## Gin's synthesis of nominine

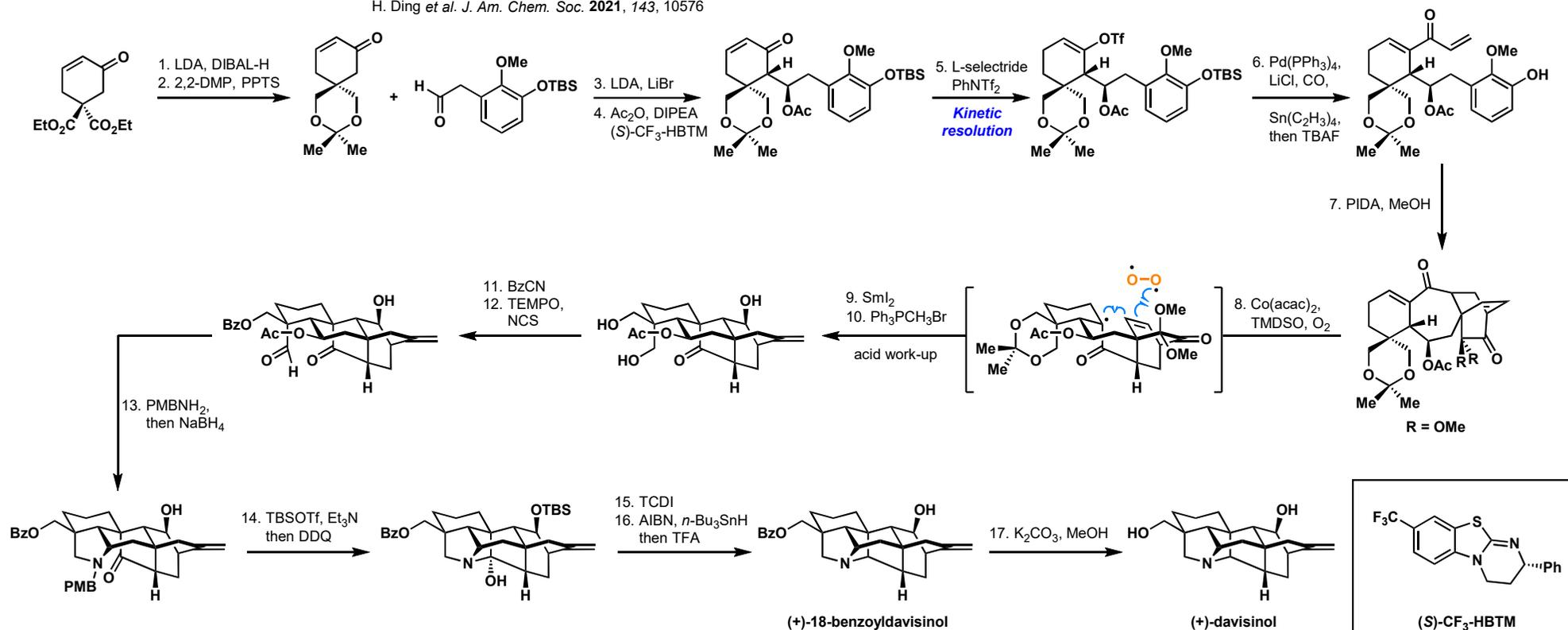
D. Y. Gin *et al.* *J. Am. Chem. Soc.* **2006**, *128*, 8734D. Y. Gin *et al.* *Chem. Eur. J.* **2008**, *14*, 1654



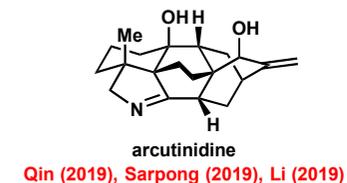
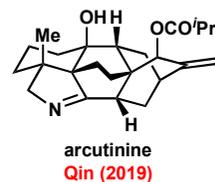
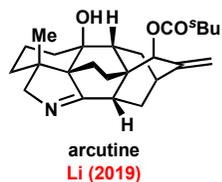
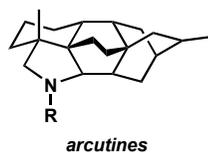
## Total Synthesis of the Hetisine-type Alkaloids



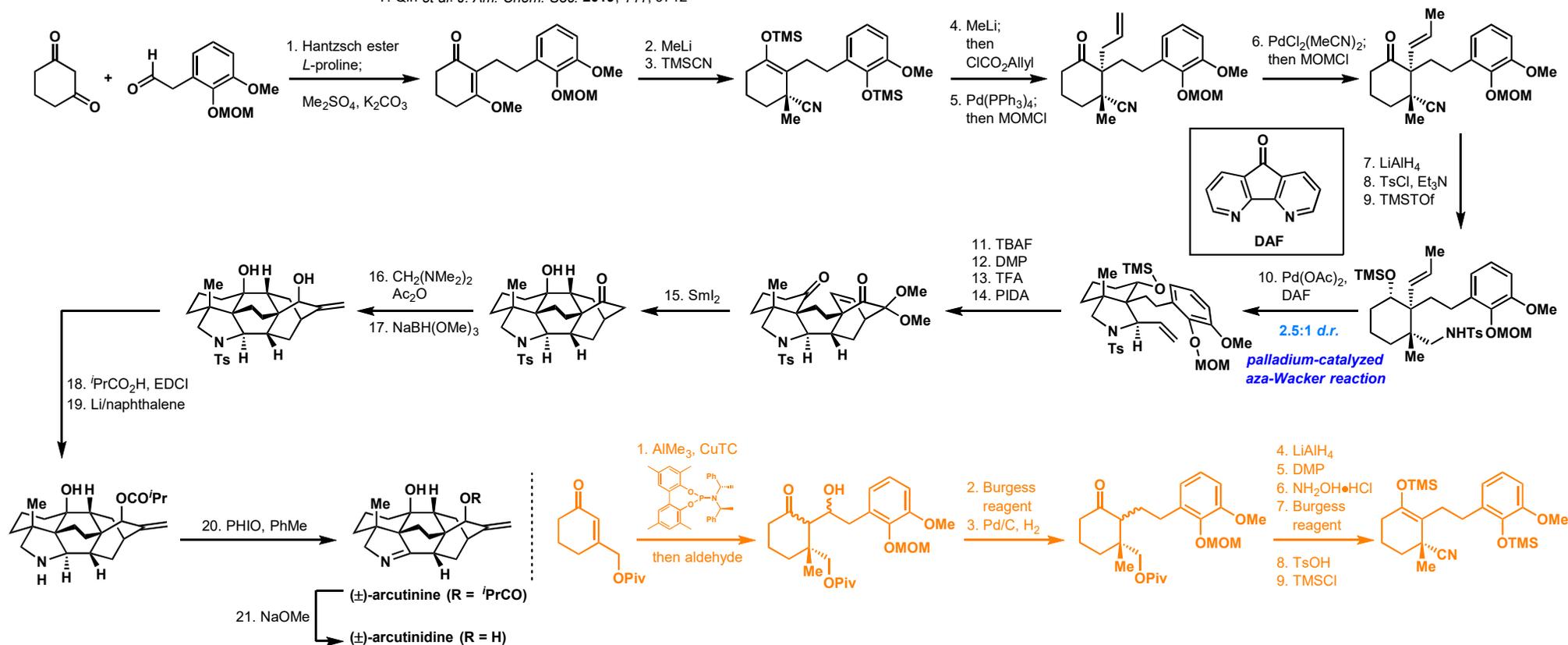
## Ding's synthesis of (+)-davisinol

H. Ding *et al.* *J. Am. Chem. Soc.* 2021, 143, 10576

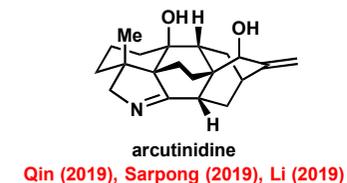
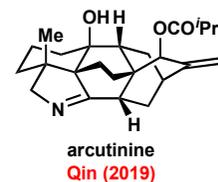
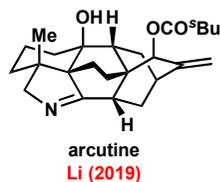
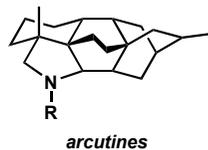
## Total Synthesis of the Hetisine-type Alkaloids



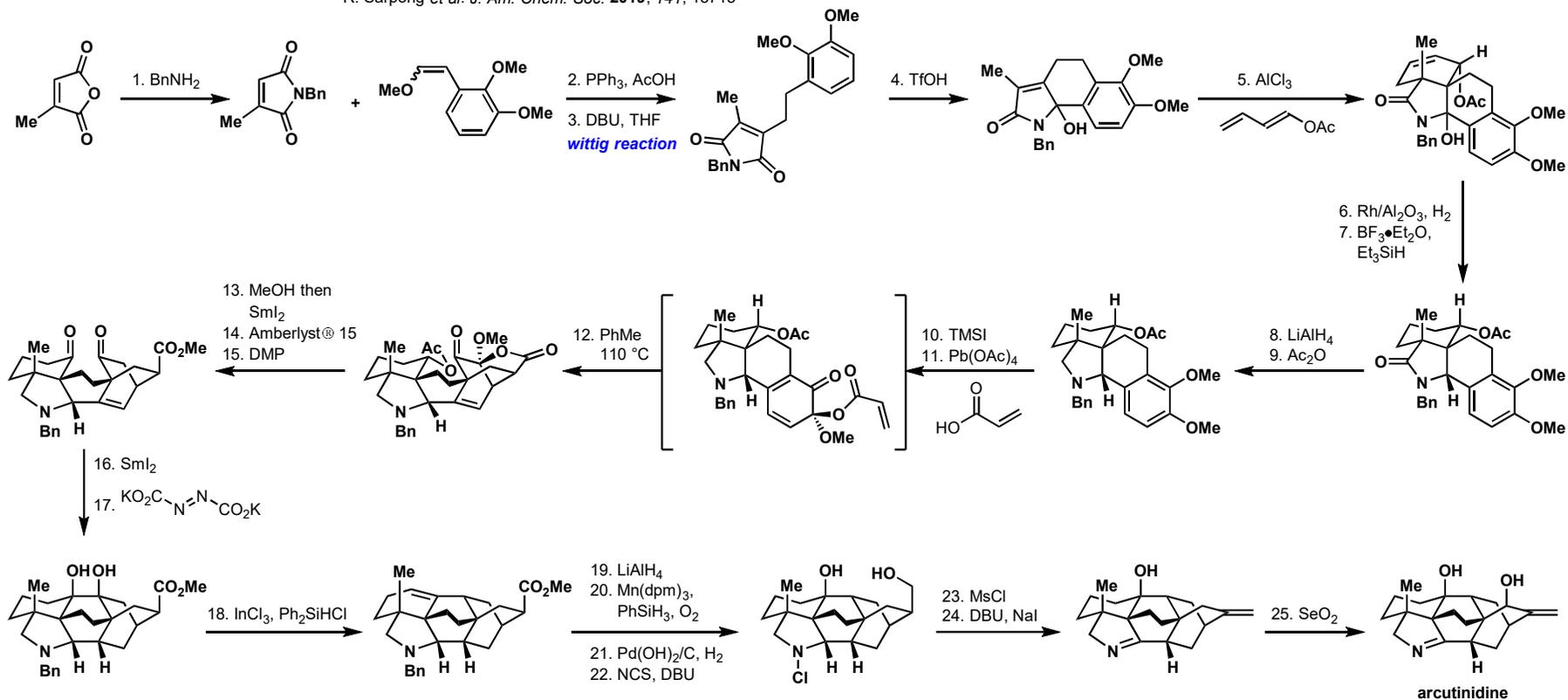
## Qin's synthesis of arcutinine and arcutinidine

Y. Qin et al. *J. Am. Chem. Soc.* 2019, 141, 9712

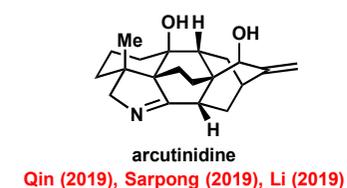
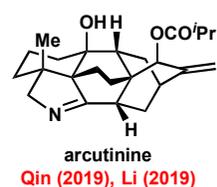
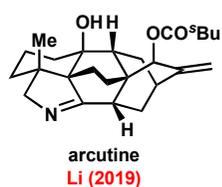
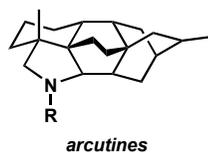
## Total Synthesis of the Hetisine-type Alkaloids



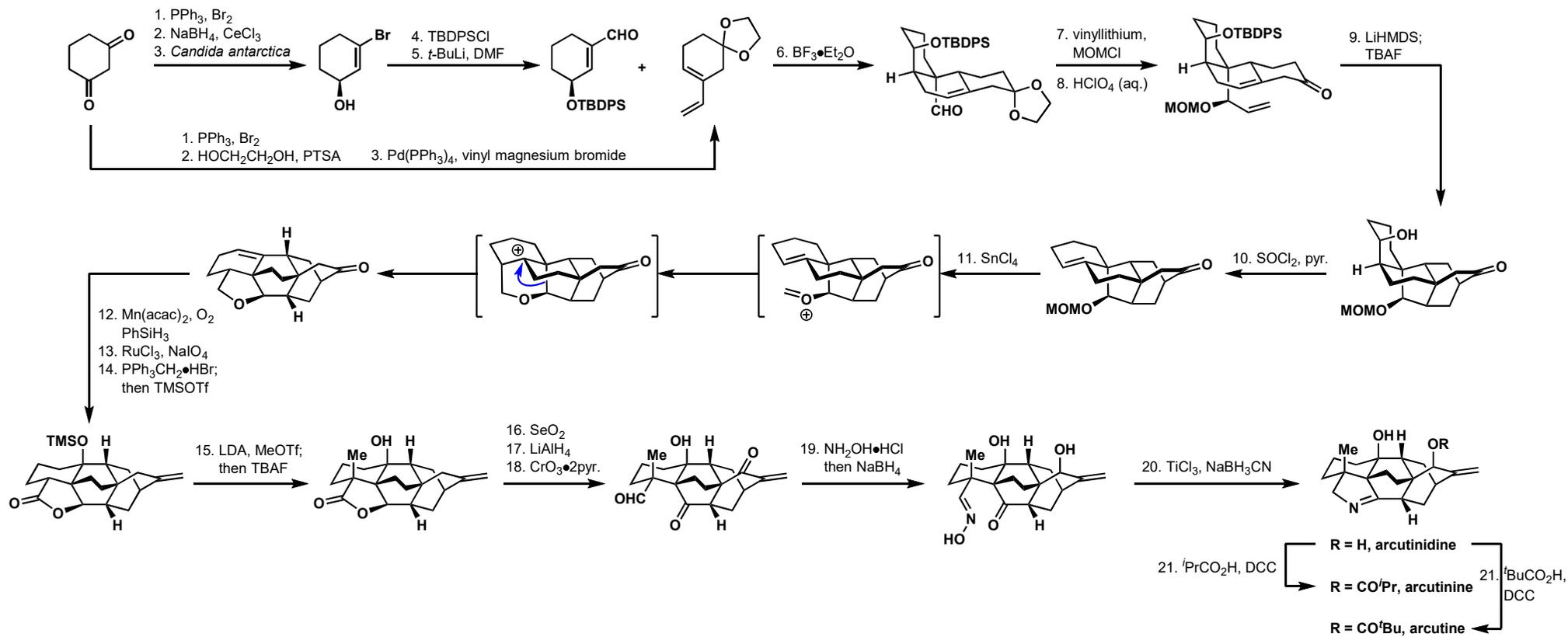
## Sarpong's synthesis of arcutinidine

R. Sarpong *et al.* *J. Am. Chem. Soc.* **2019**, *141*, 13713

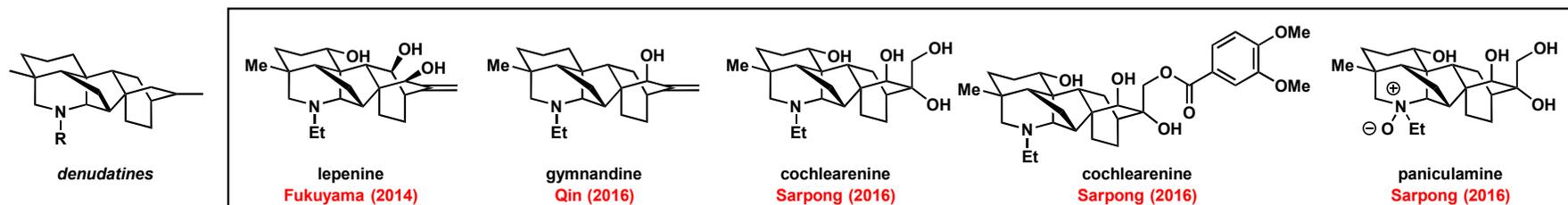
## Total Synthesis of the Hetisine-type Alkaloids



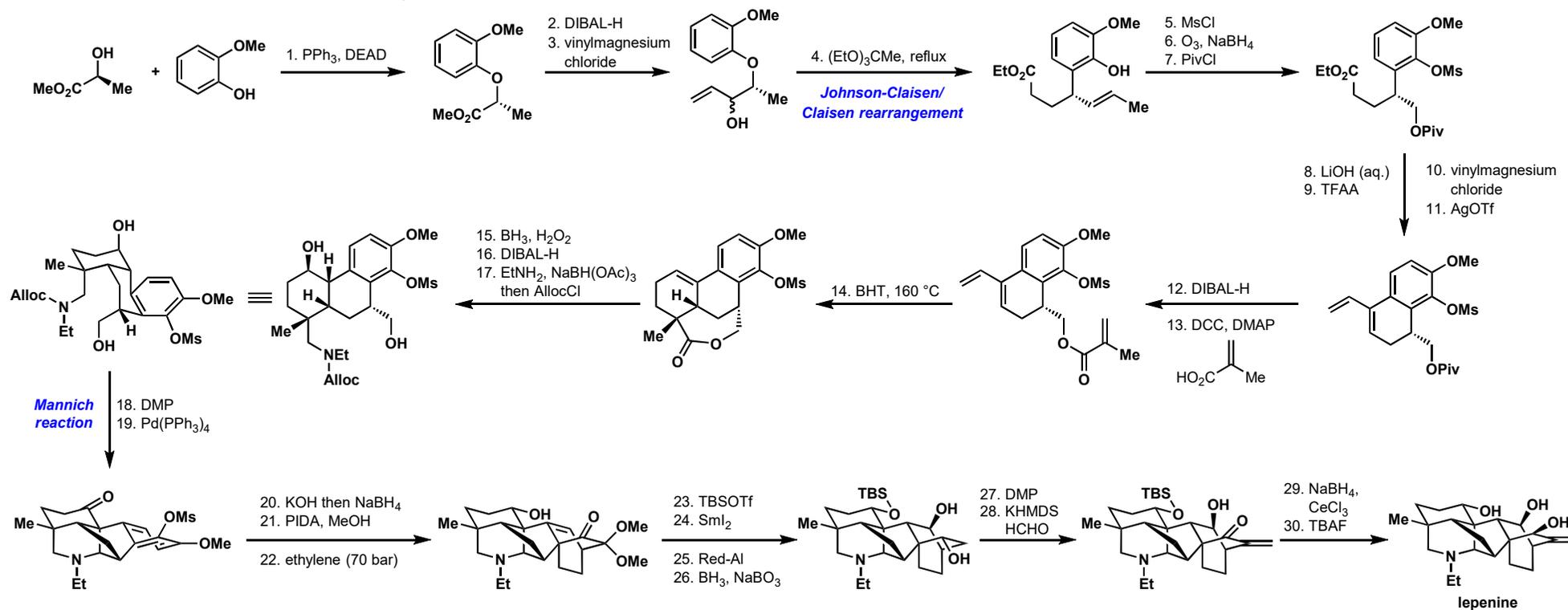
## Li's synthesis of arcutine, arcutinine, arcutinidine

A. Li et al. *J. Am. Chem. Soc.* **2019**, *141*, 13718

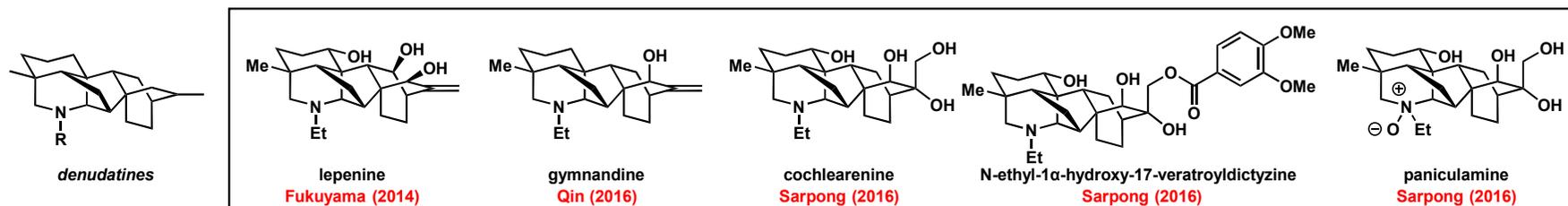
## Total Synthesis of the denudatine-type Alkaloids



## Fukuyama's Synthesis of the lepenine

T. Fukuyama *et al.* *J. Am. Chem. Soc.* **2014**, *136*, 6598

## Total Synthesis of the denudatine-type Alkaloids



## Sarpong's Synthesis of the denudatine-type alkaloids

R. Sarpong *et al.* *J. Am. Chem. Soc.* **2016**, *138*, 10830