

Moreno, J.; Picazo, E.; Morrill, L. A.; Smith, J. M.; Garg, N. K. J. Am. Chem. Soc. 2016, 138, 1162.



## 4 Subclasses of Akuammiline Natural Products



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**2** 2/13/2016

# **Biological Activity**

• Over 30 alkaloids isolated from the plant *Alstonia scholaris*.



- Many of these alkaloids display anti-tussive, anti-asthmatic and anti-inflammatory effects
- Some exhibit opioid behavior and cytotoxic activity

#### **Retrosynthetic Analysis**



#### Forward Synthesis to Au Cyclization



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#### Au-Mediated Cyclization



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Mike Hoursh Ring Wijhang Up; Zhou, Y.; Zhao, L.; Jiang, H.; Liu, HPage & Gham. 2010, 75, 5810. 2. Lu, Z.; Li, Y.; Deng, J.; Li, A. Nat. Chem. 2013 2016 79.

## Forward Synthesis to Interrupted Fischer Indole



# Interrupted Fischer Indole



## Forward Synthesis to the Methanoquinozilidines



Maligres, P. E.; Houpis, I.; Rossen, K.; Molina, A.; Sager, J.; Upadhyay, V.; Wells, K. M.; Reamer, R. A.; Lynch, J. E.; Askin, D.; 2/13/2016 Volante, R. P.; Reider, P. J.; Houghton, P. *Tetrahedron* **1997**, 53, 10983.

### Completion the Methanoquinozilidines



# Conclusions

 Successfully completed first enantioselective route to (–)-Aspidophylline A

• They were able to synthesize two natural products from the methanoquinolizidine akuammilines, for which there was no previous synthesis.