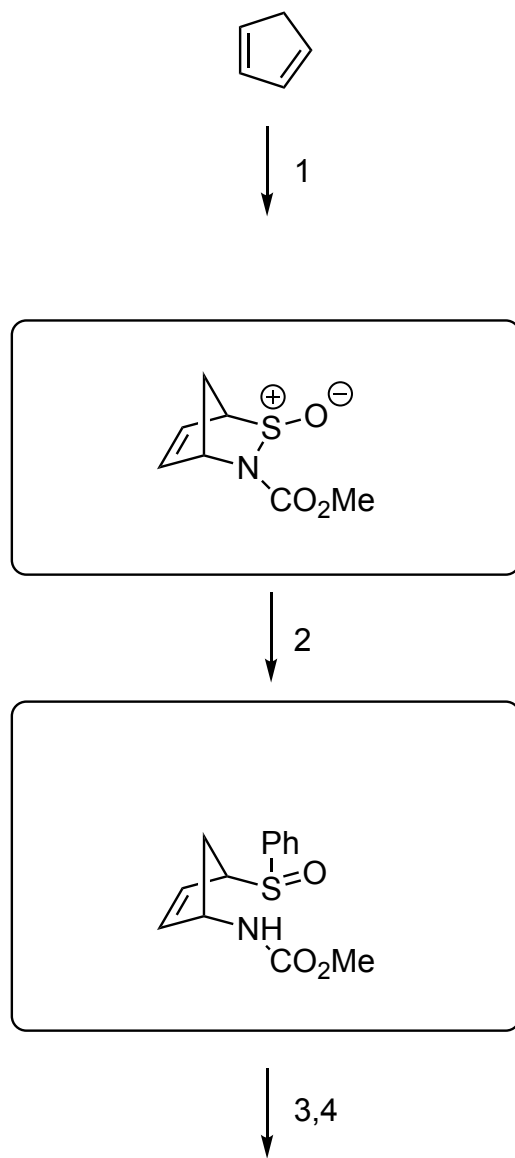


Studies on Total Synthesis of the Cytotoxic Marine Alkaloid Agelastatin A

Anderson, G. T.; Chase, C. E.; Koh, Y.-H.; Stien, D.; Weinreb, S. M.

J. Org. Chem. **1998**, *63*, 7594

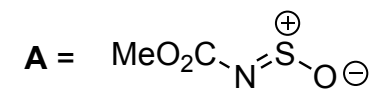


1) **A**, Benzene, 0 °C

2) PhMgBr, THF, -60 °C

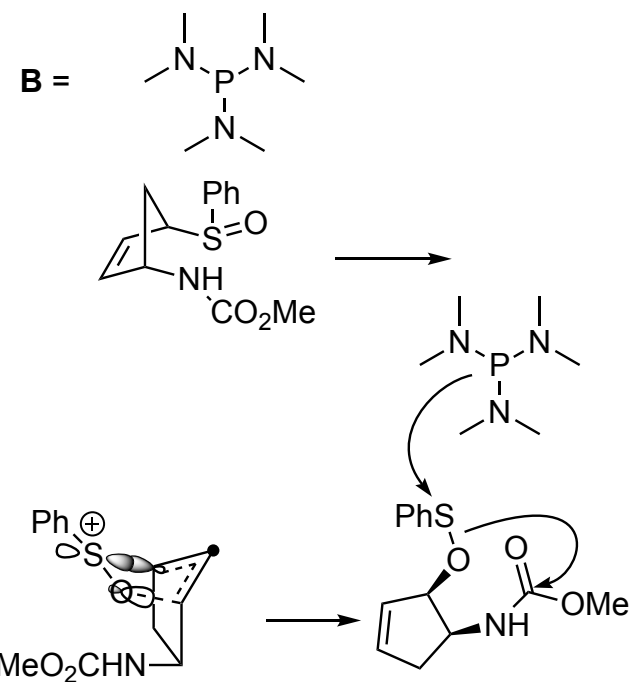
3) **B**, EtOH, 80 °C

4) Boc₂O, DMAP, Et₃N, MeCN, 80 °C



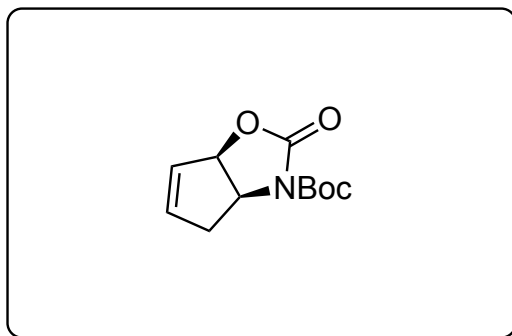
How would you make **A**?

O-Methylcarbamate + SOCl₂

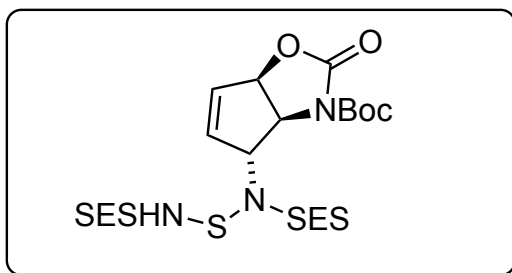


What is the name of this reaction? *Mislow-Evans*

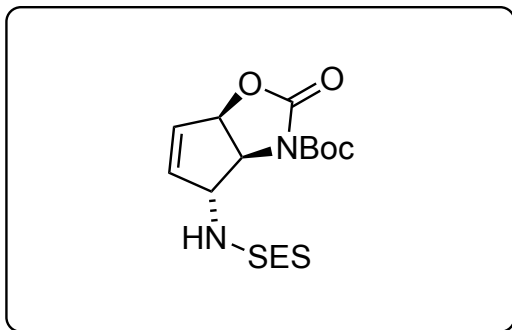
Please draw a mechanism and the frontier molecular orbitals.



5



6

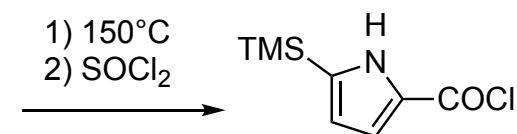
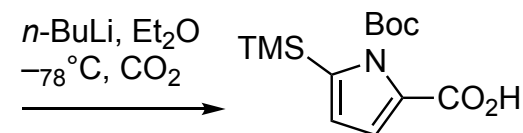
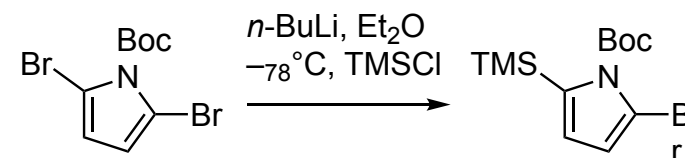
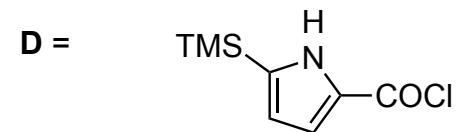
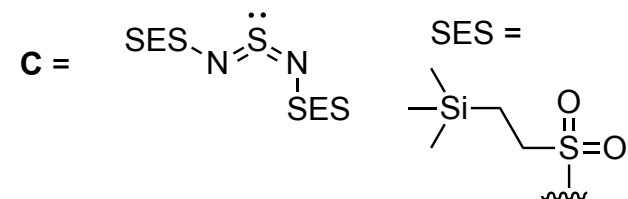


7

5) **C**, Benzene, 80 °C

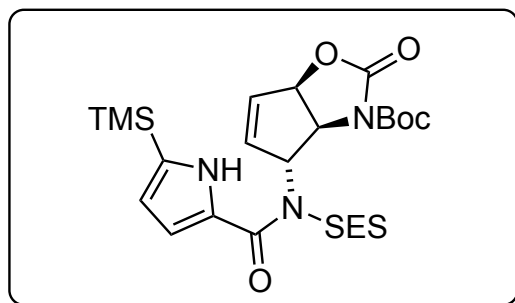
6) NaBH₄ or P(OMe)₃, MeOH

7) **D**, DMAP, Et₃N, THF

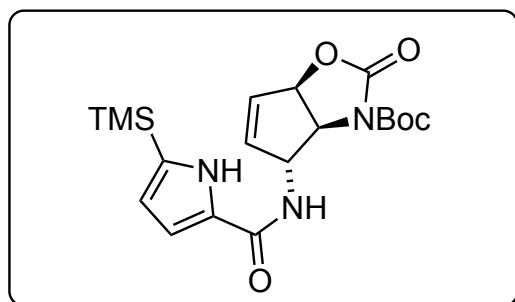


NOTE: If they say TFA for deprotection of Boc-group, question them on why this might be bad (Protodesilylation).

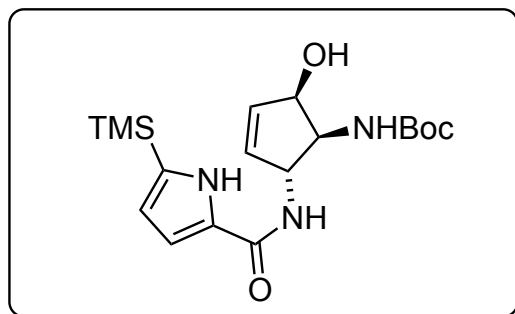
*Provide a plausible synthesis for **D** starting with pyrrole.*



↓ 8



↓ 9

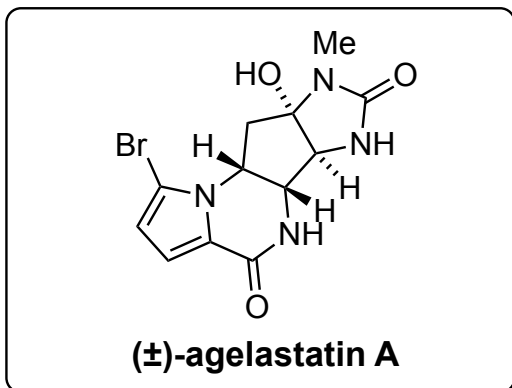
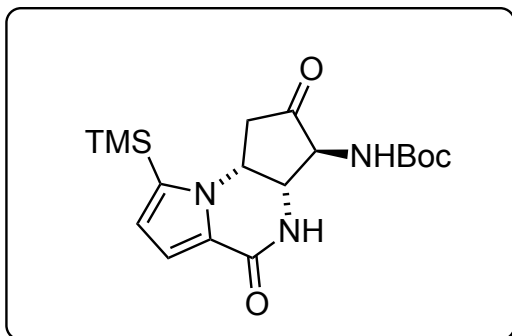


↓ 10,11

8) TBAF, THF

9) LiOH, THF, Water

10) PDC, DMF
11) Cs₂CO₃, MeOH



- 12) NBS, THF
13) TMSI, DCM
14) MeNCO, NaOH, H₂O