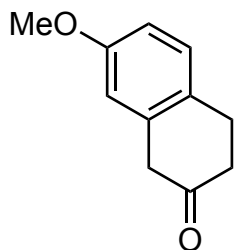


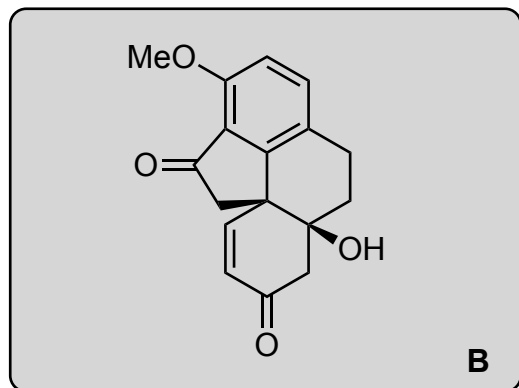
## Total Synthesis of (-)-Morphine

Umihara, H.; Yokoshima, S.; Inoue, M.; Fukuyama, T.

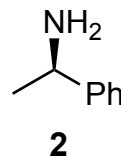
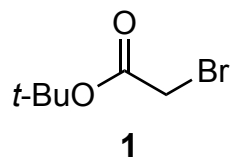
*Chem. Eur. J.* **2017**, *23*, 6993–6995



1-6



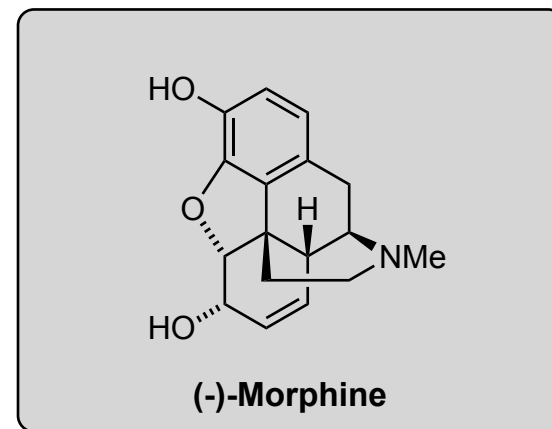
- 1) pyrrolidine, reflux *then* **1**, *t*-BuOH *then* AcOH buffer
- 2) **2**, reflux *then* methyl vinyl ketone *then* AcOH buffer
- 3) TfOH
- 4) PhSeCl, MeCN/H<sub>2</sub>O (5:1)
- 5) *m*CPBA, NaHCO<sub>3</sub>
- 6) Cs<sub>2</sub>CO<sub>3</sub>, 60 °C



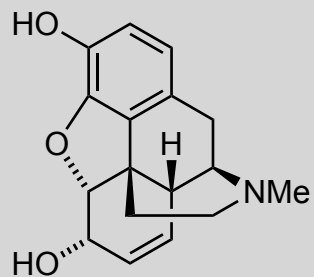
3) Name of reaction?

Metal-free Friedel-Crafts

6) Name or reactions? *hint*: same reaction but different directions  
retro-aldol then aldol

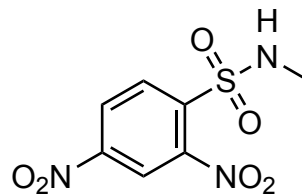


7-18



**(-)-Morphine**

- 7) *m*CPBA, NaHCO<sub>3</sub>
- 8) K<sub>2</sub>CO<sub>3</sub>, MeOH then TMSCHN<sub>2</sub>
- 9) NaBH<sub>4</sub>
- 10) TsOH•H<sub>2</sub>O
- 11) LiAlH<sub>4</sub>
- 12) **3**, DEAD, Ph<sub>3</sub>P
- 13) O<sub>2</sub>, *hν* (405 nm), tetraphenylporphine  
*then* Et<sub>3</sub>N
- 14) Martin's sulfurane
- 15) PhSH, iPr<sub>2</sub>NEt
- 16) 4M HCl in dioxane
- 17) NaBH<sub>4</sub>
- 18) BBr<sub>3</sub>



**3**

7) Name of the reaction?

Baeyer-Villiger reaction

12) Name or reaction

Mitsunobu reaction

13) *hint*: Et<sub>3</sub>N promotes O-O bond cleavage.

15) *hint*: protecting group cleavage

17) name of natural product?

Codeine