

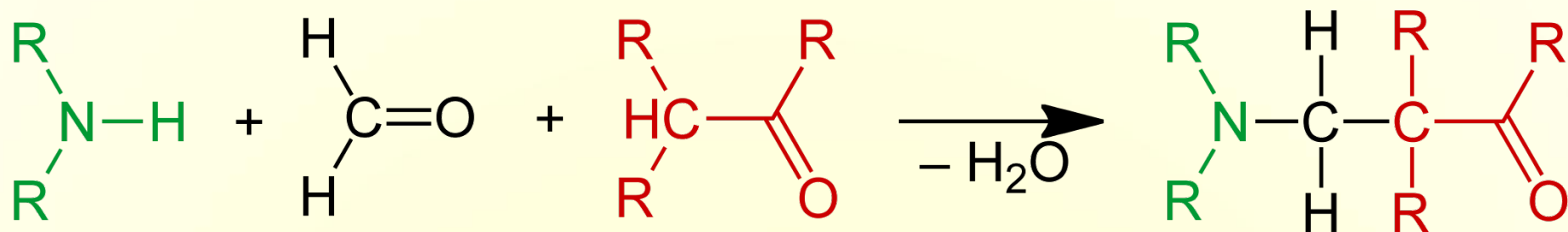
MANNICH REACTION

2016-5-10

WZQ

MANNICH REACTION

Mannich reaction

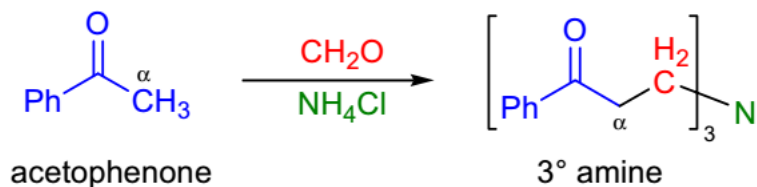


The Mannich reaction is an organic reaction which consists of an amino alkylation of an acidic proton placed next to a **carbonyl functional group** by **formaldehyde** and a **primary or secondary amine or ammonia**.

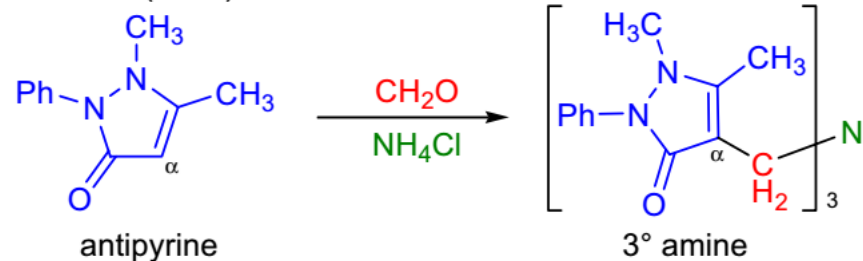
The final product is a β -amino-carbonyl compound also known as a Mannich base

MANNICH REACTION

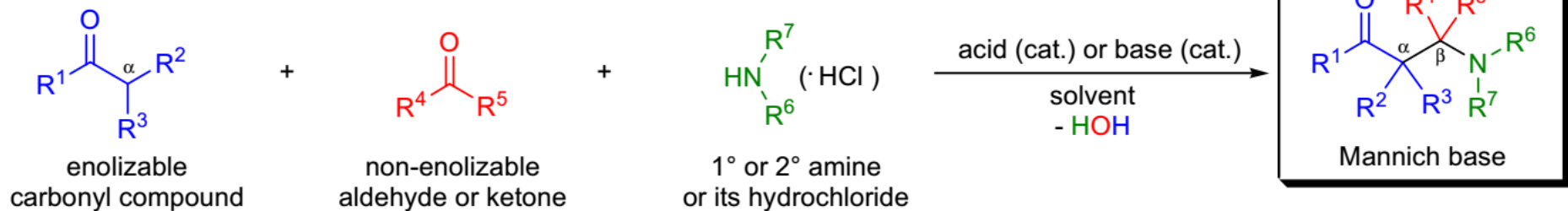
Tollens and von Marle (1903):



Mannich (1917):



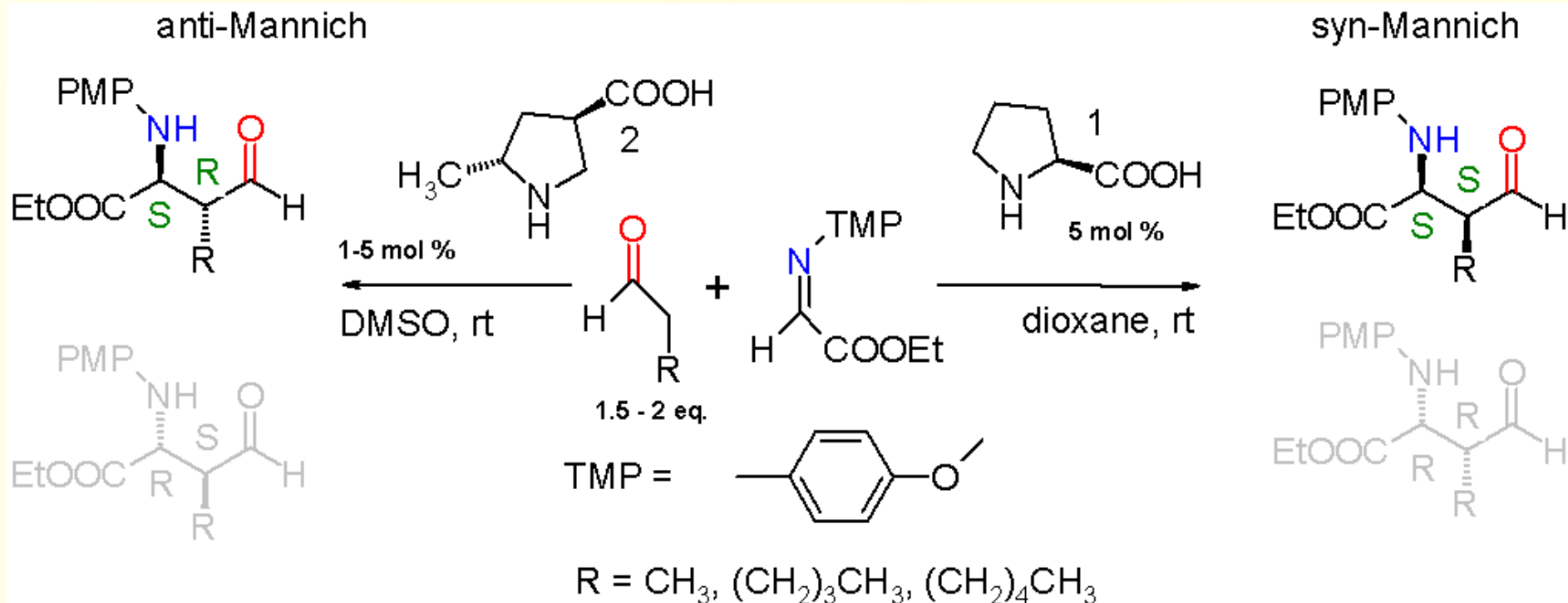
Mannich reaction:



$\text{R}^1 = \text{H, alkyl, aryl, OR}; \text{R}^{2-3} = \text{H, alkyl, aryl}; \text{R}^{4-5} = \text{H, alkyl, aryl}; \text{R}^6 = \text{H, alkyl, OH, NH}_2; \text{R}^7 = \text{H, alkyl}; \text{solvent} = \text{ROH, H}_2\text{O, AcOH}$

MANNICH REACTION

Asymmetric Mannich reactions

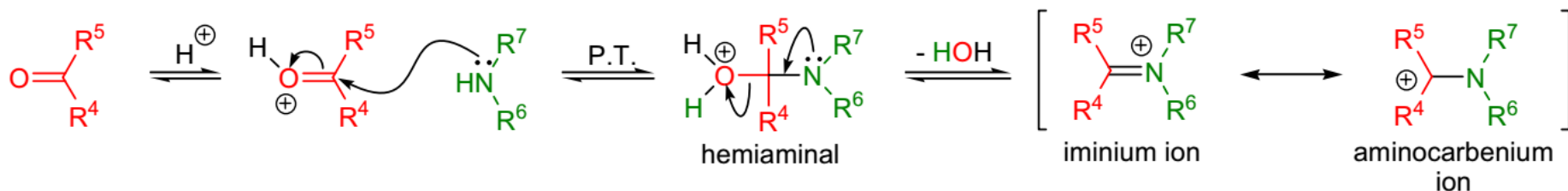


Journal of the American Chemical Society **124** (9): 1866–1867

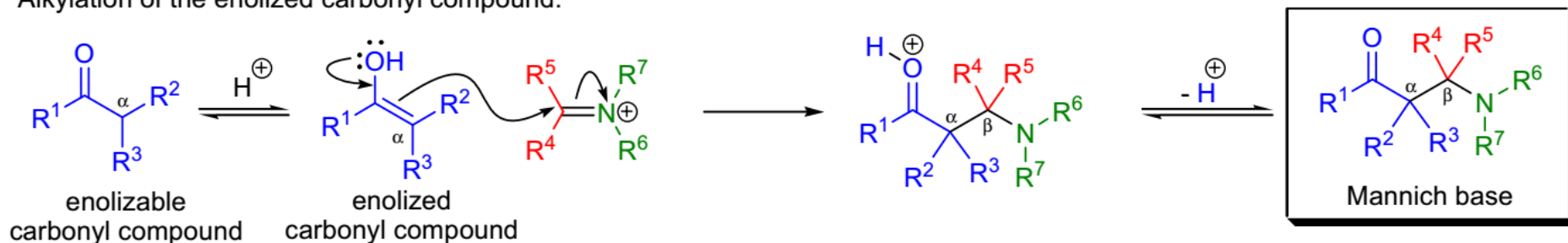
MANNICH REACTION

Mechanism:

Formation of the reactive iminium ion under acidic conditions:



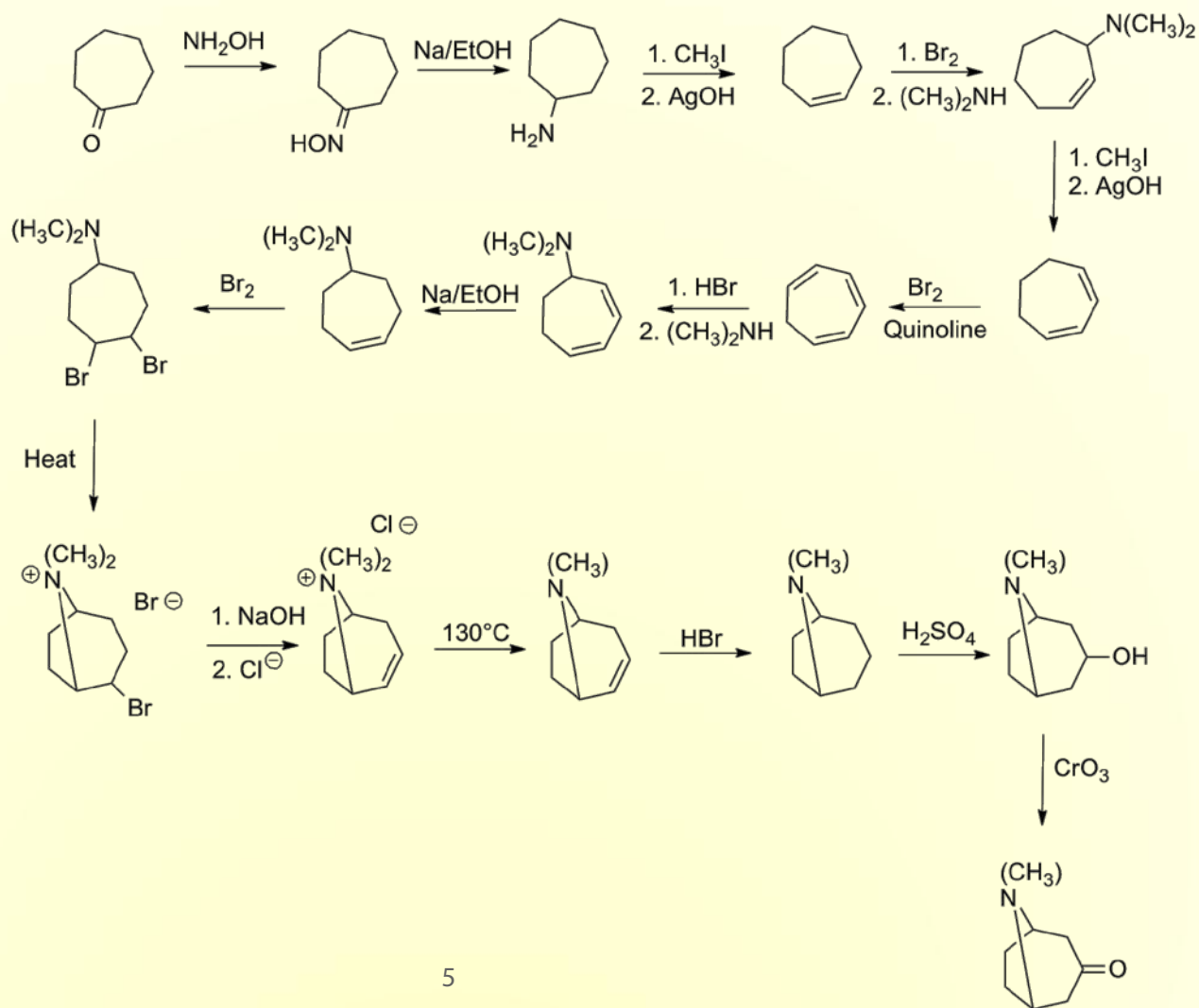
Alkylation of the enolized carbonyl compound:



Generally it is the addition of resonance-stabilized carbon nucleophiles to iminium salts and imines

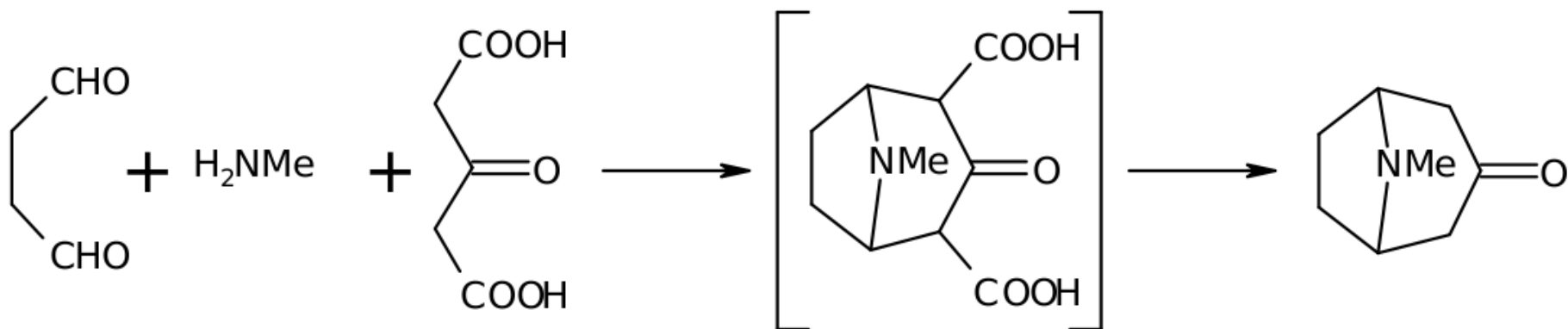
MANNICH REACTION

Applications: Tropinone

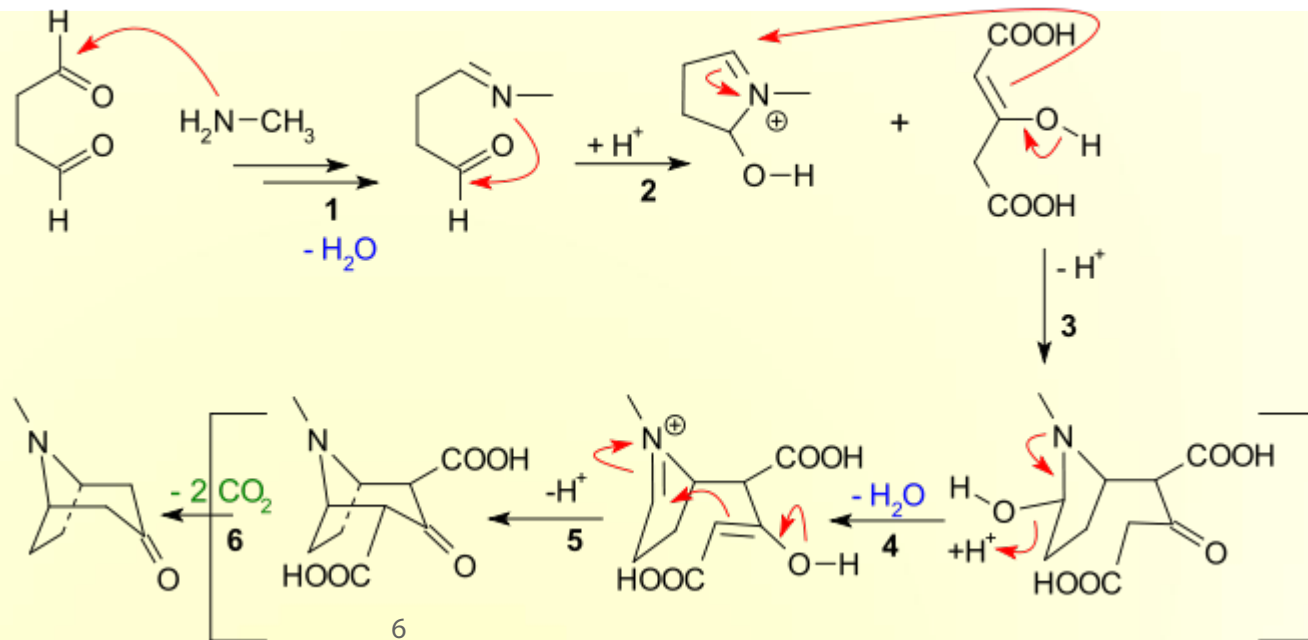


MANNICH REACTION

Applications: Tropinone

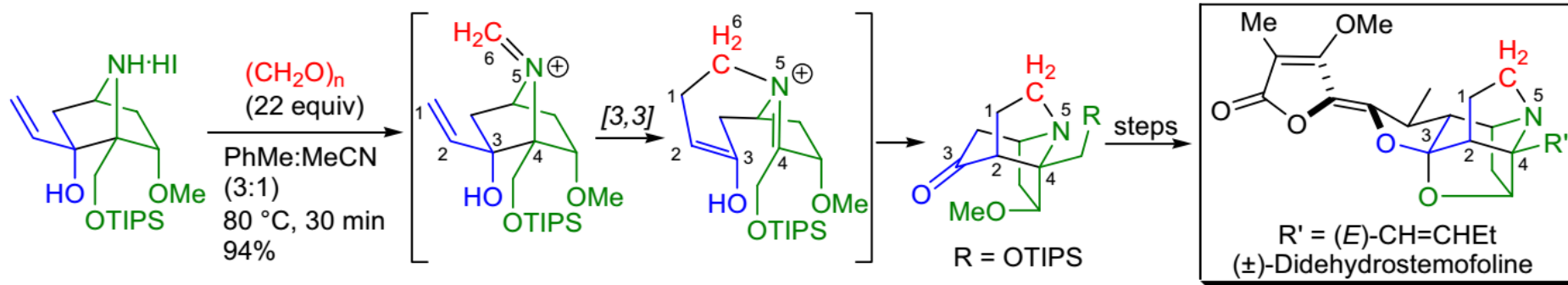
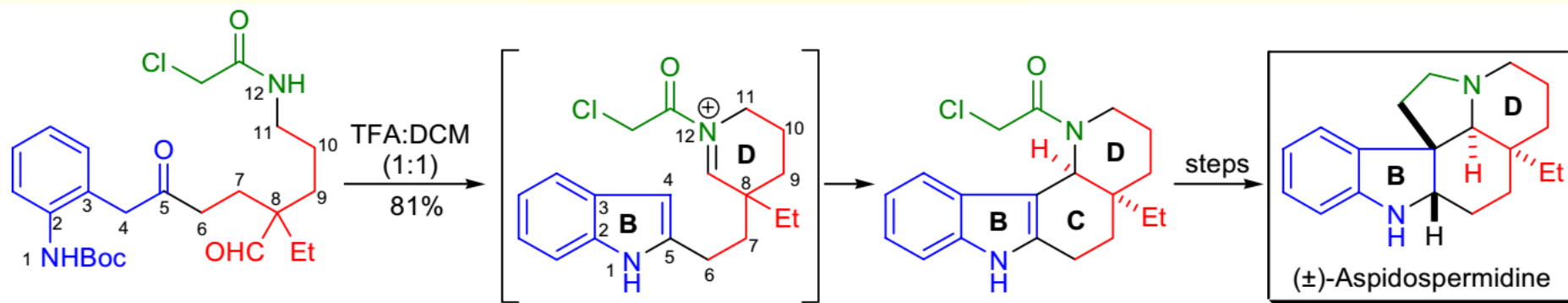


Mechanism:



MANNICH REACTION

Applications:



MANNICH REACTION

Applications:

