



Collective synthesis of natural products by means of organocascade catalysis

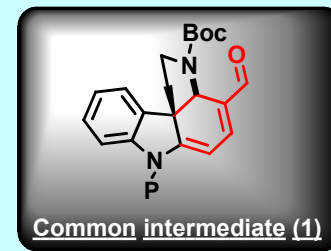
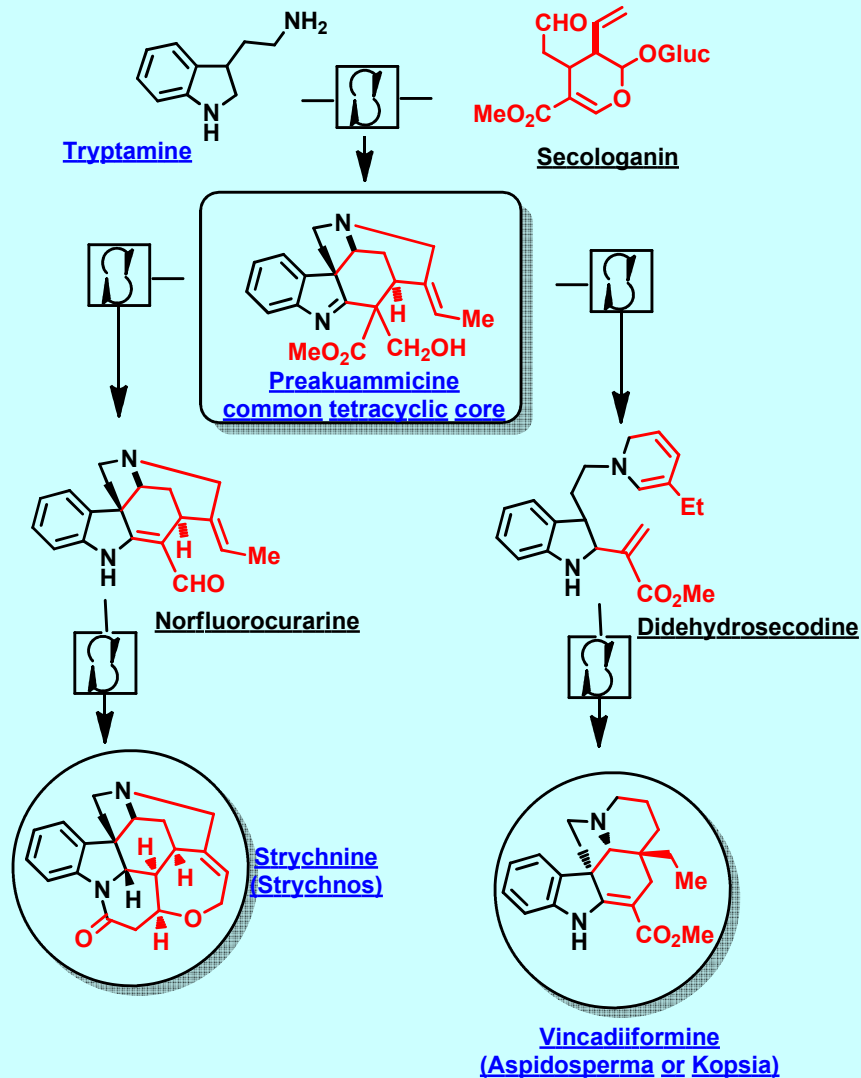
Spencer B. Jones, Bryon Simmons, Anthony Mastracchio &
David W. C. MacMillan*

NATURE 475, 183, 2011

Princeton University
USA

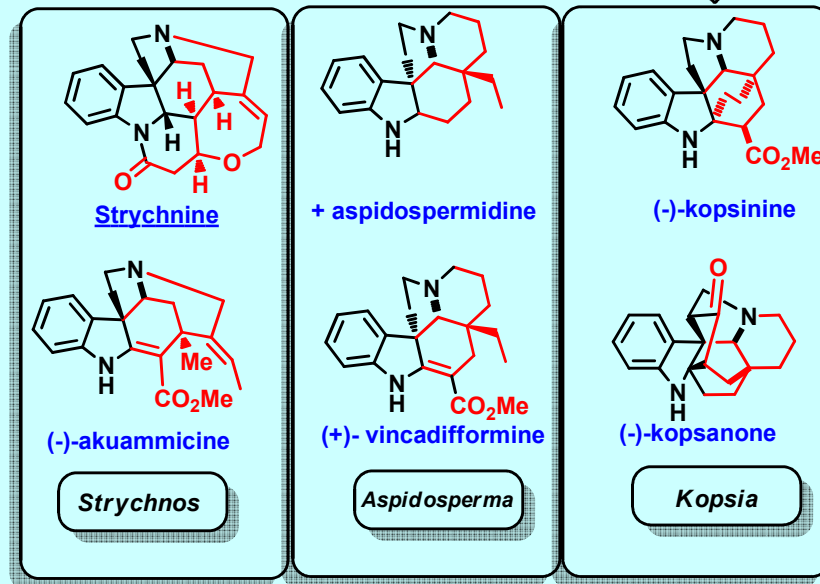
Merck Center for Catalysis at Princeton University, New Jersey USA

Introduction.....

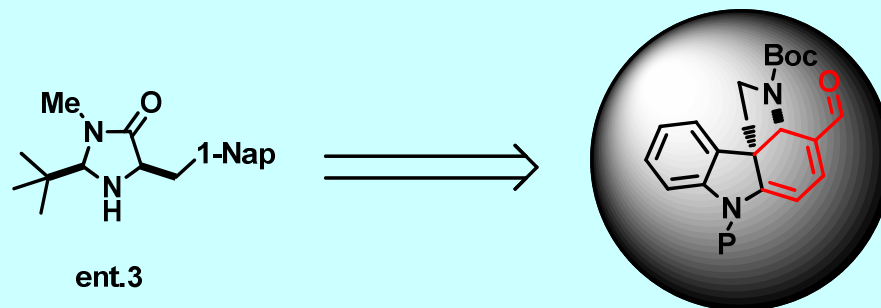
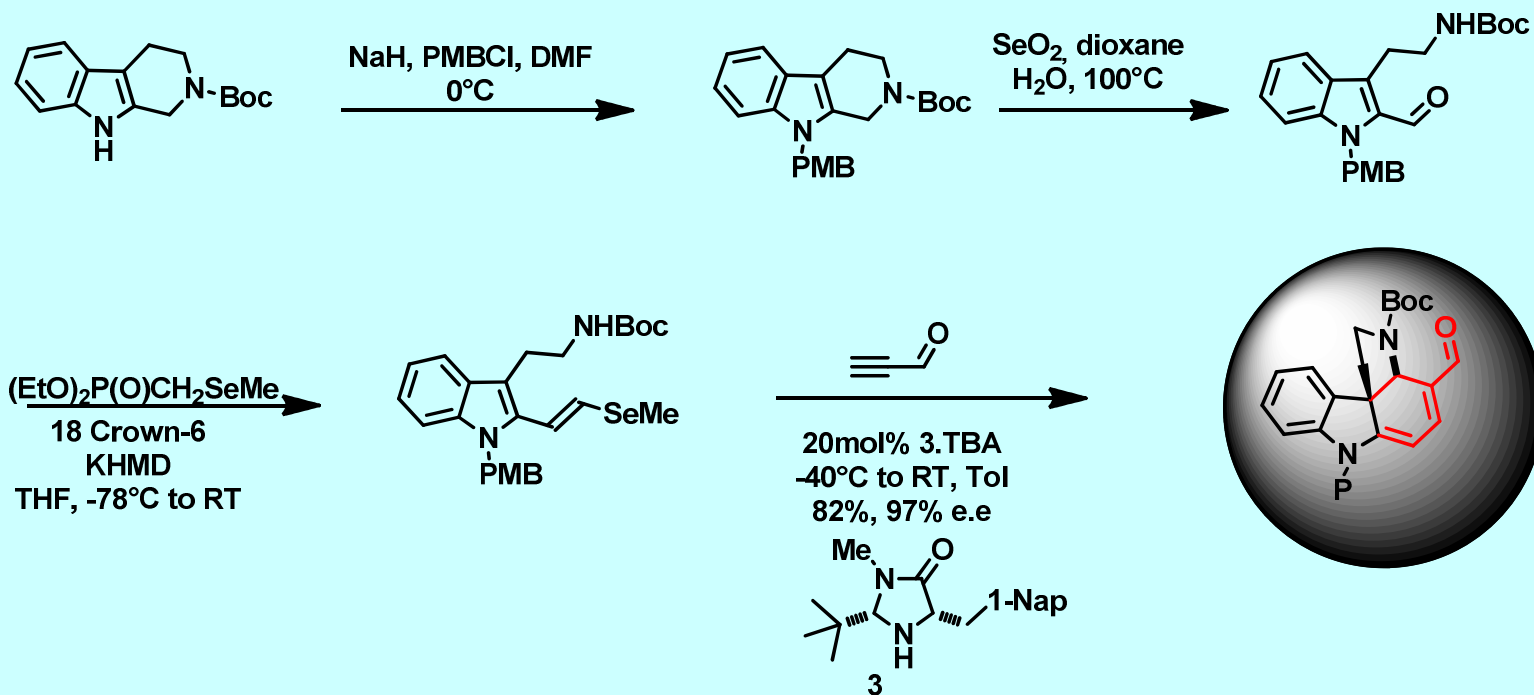


Synthetic elaboration

Enantioselective synthesis of Strychnos, Aspidosperma and Kopsia alkaloids

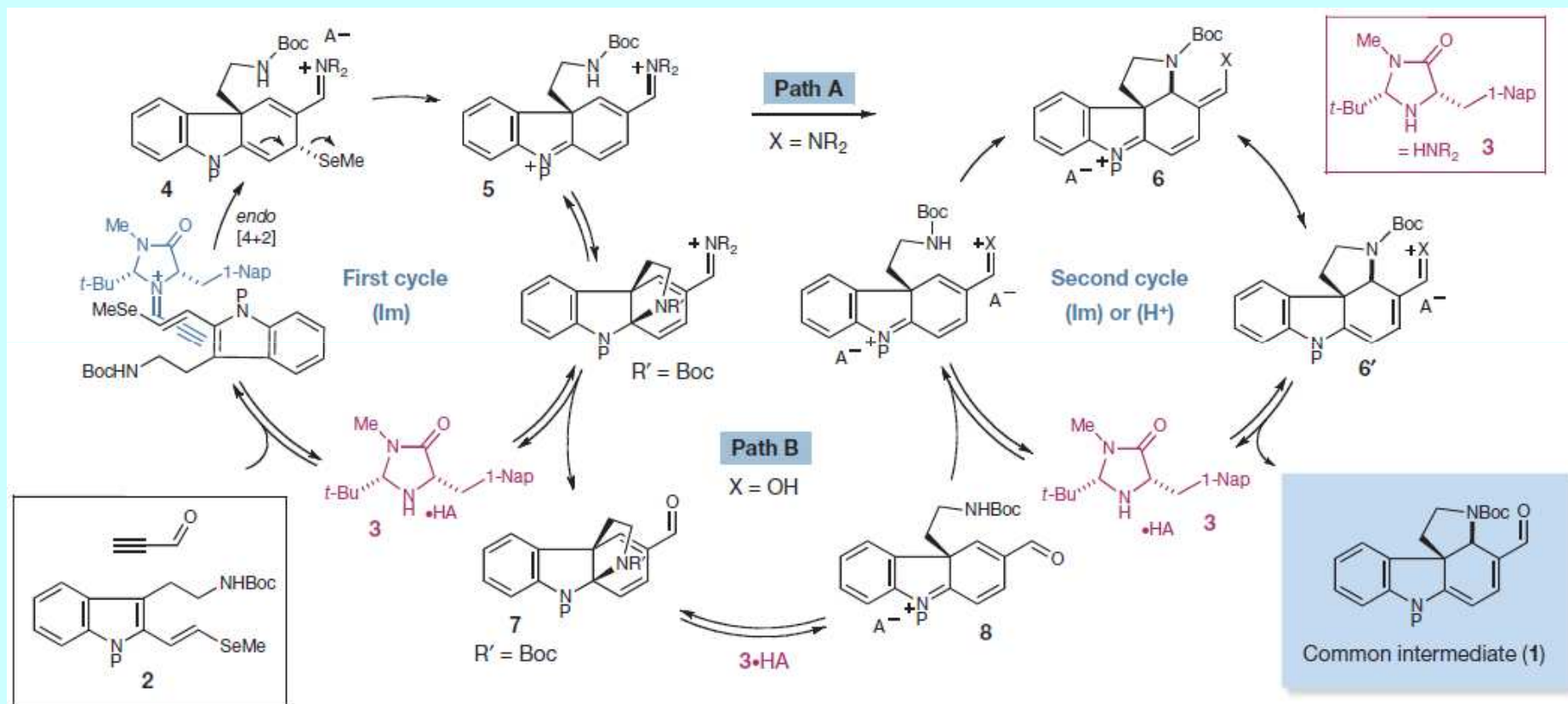


Synthesis of Common Intermediate

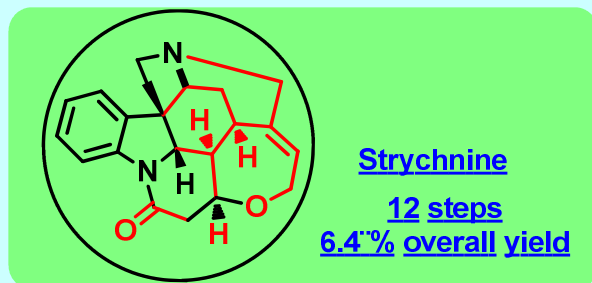
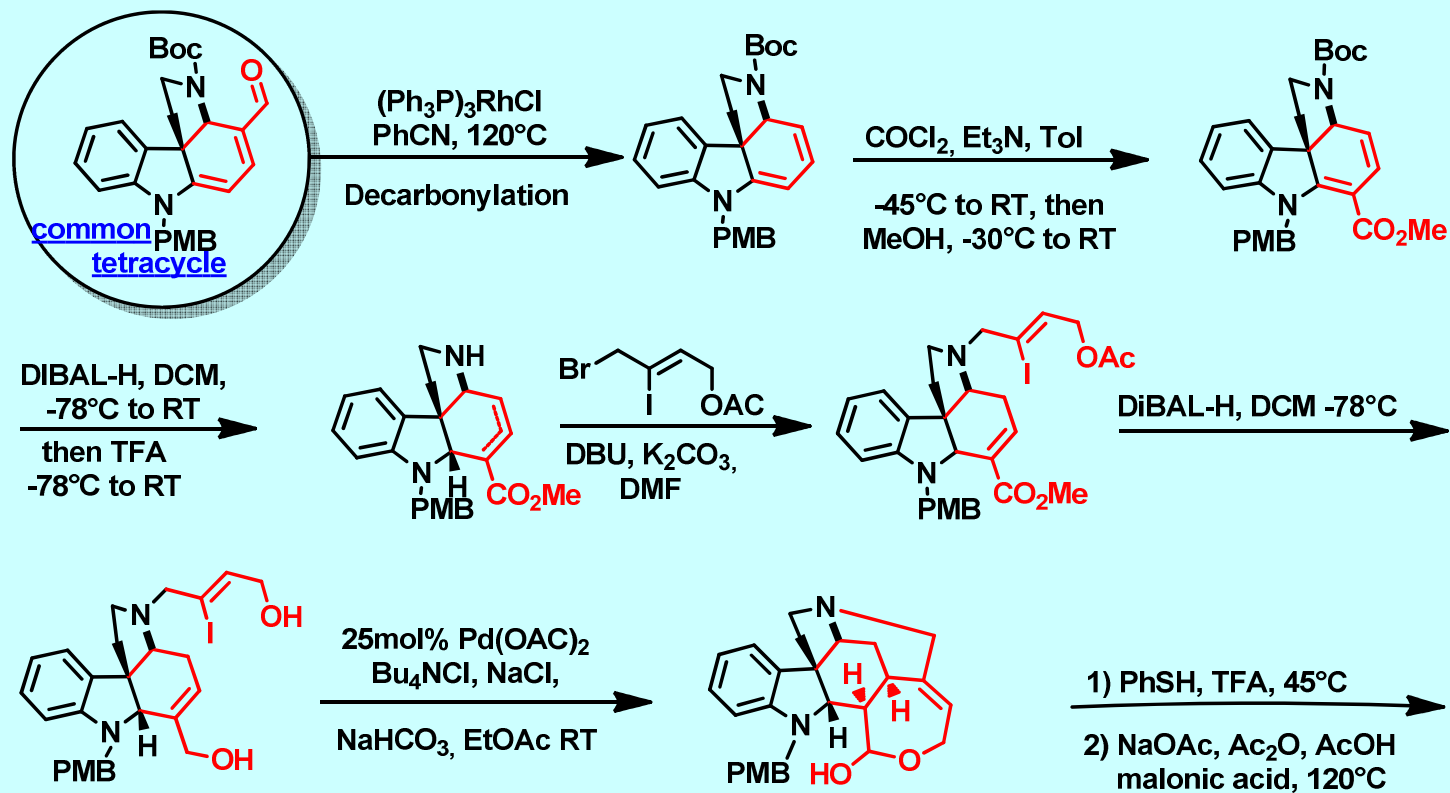


**common
intermediate**

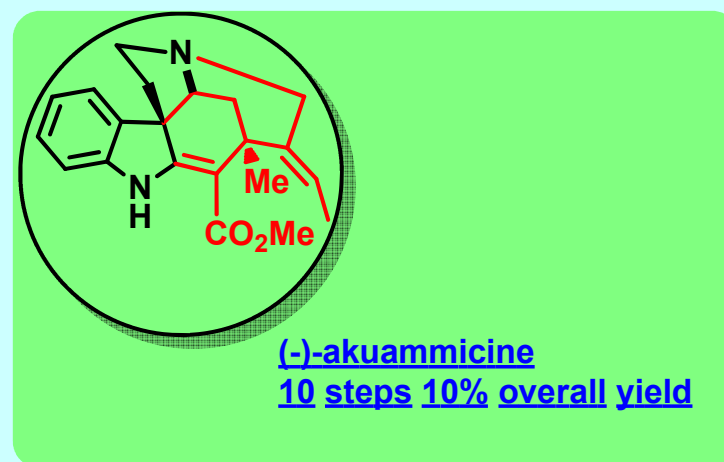
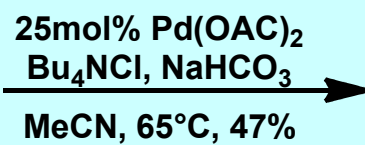
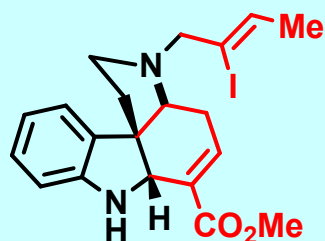
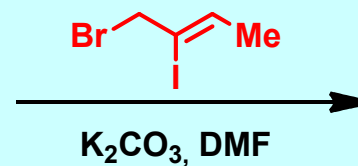
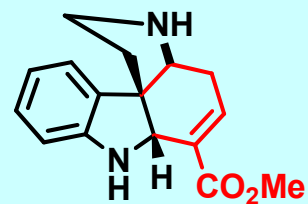
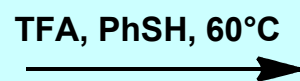
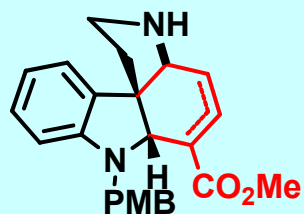
Proposed mechanism of organocascade cycles for the generation of common tetracyclic intermediate



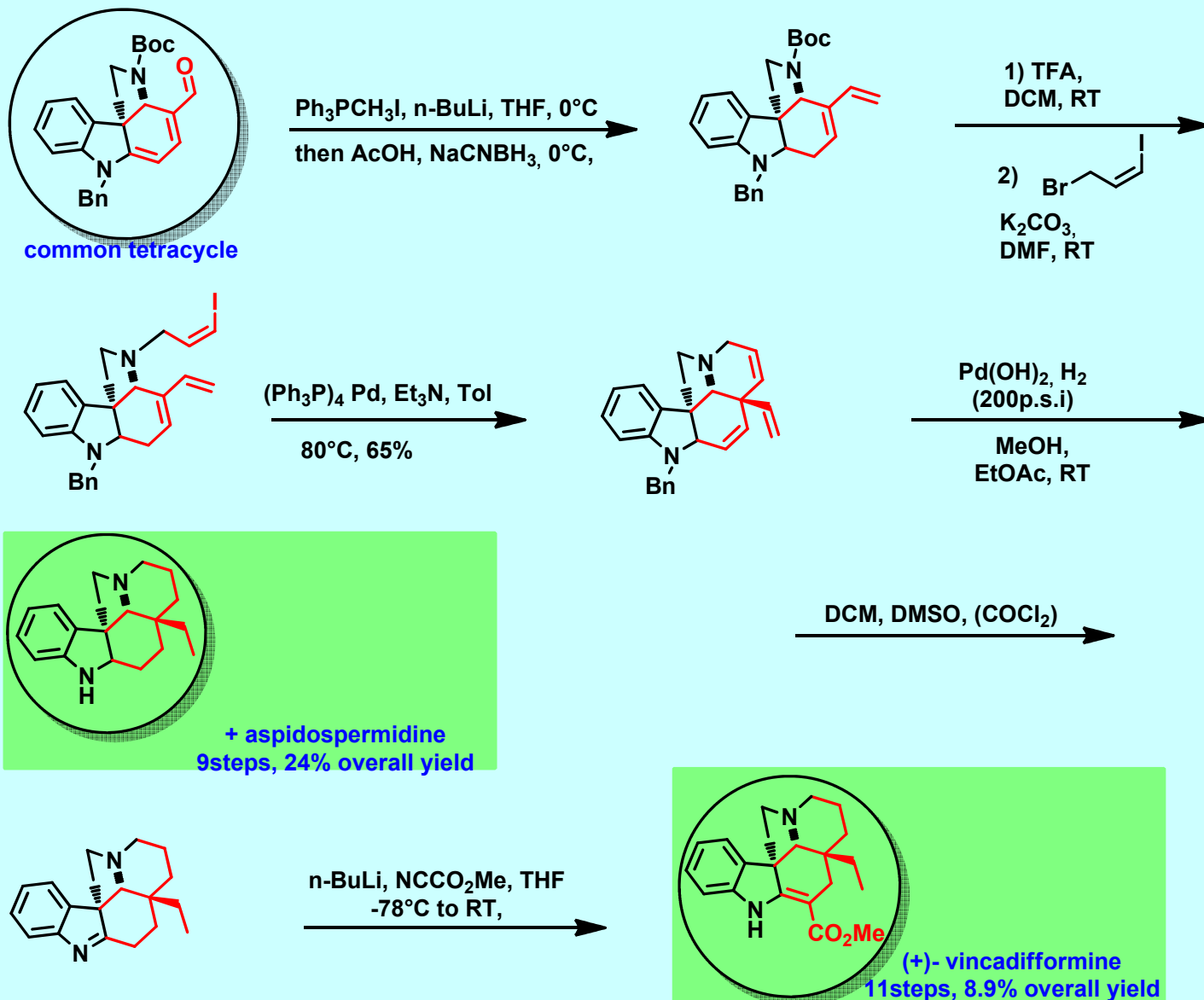
Synthesis of Strychnine



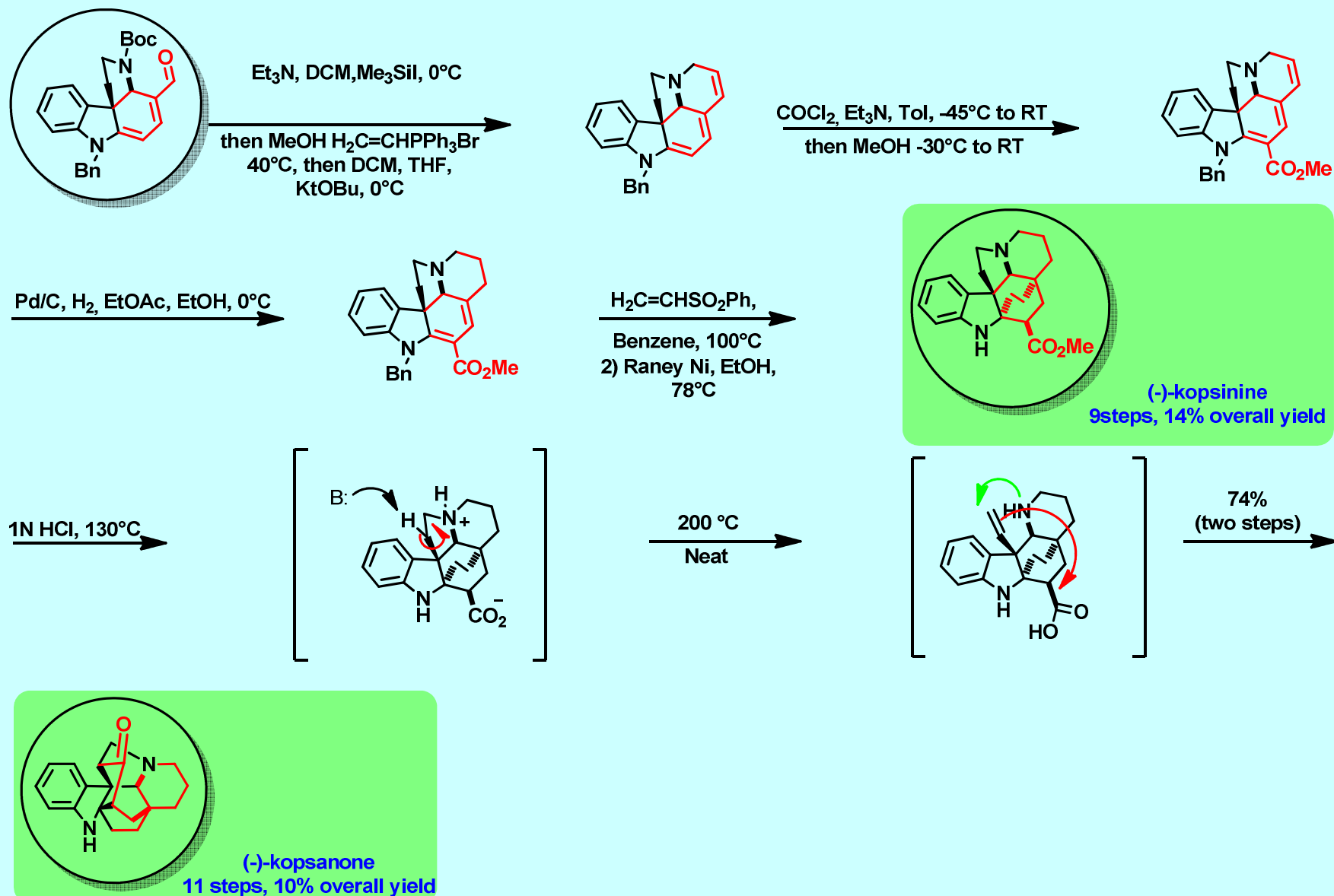
Synthesis of Akuammicine



Synthesis of Aspidospermidine & Vincadiiformine

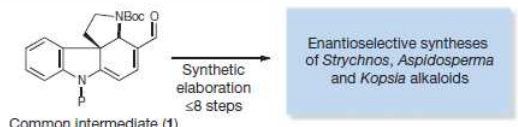


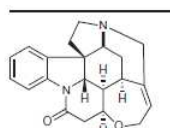
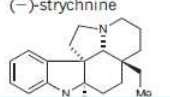
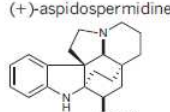

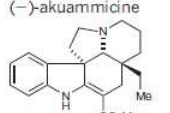
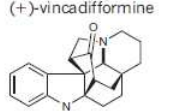
Synthesis of Kopsanone & Kopsinine



conclusion

Table 1 | Enantioselective synthesis of six well-known indole alkaloids



Compound	No. steps here*	Overall yield (%)	PSAC steps	PSCA steps
 (-)-strychnine	12	6.4	25 (refs 19, 20)	16 (ref. 21)
 (+)-aspidospermidine	9	24	13 (ref. 30)	11 (ref. 29)
 (-)-kopsinine	9	14	NA	19 (ref. 32)
 (-)-akuammicine	10	10	NA	NA
 (+)-vincadifformine	11	8.9	NA	10 (ref. 31)
 (-)-kopsanone	11	10	NA	NA

Step counts represent the longest linear sequence from commercially available 9. NA, not applicable. PSAC, previous shortest asymmetric catalytic synthesis; PSCA, previous shortest chiral auxiliary or chiral pool synthesis.

➤ **Novel Asymmetric approach to Total synthesis based on the application of these two nature-inspired concepts, namely collective total synthesis and organocascade catalysis.**

➤ **Shortest Asymmetric synthesis of (-)-Strychnine.**

➤ **Hope in terms of natural product and medicinal agent families in the near future.**

➤ **These collective asymmetric syntheses took a total of 34 steps for 6 Natural products (Previous 76 steps)**

**Merci pour votre
attention**

photo
Sawat Pakistan

nature