

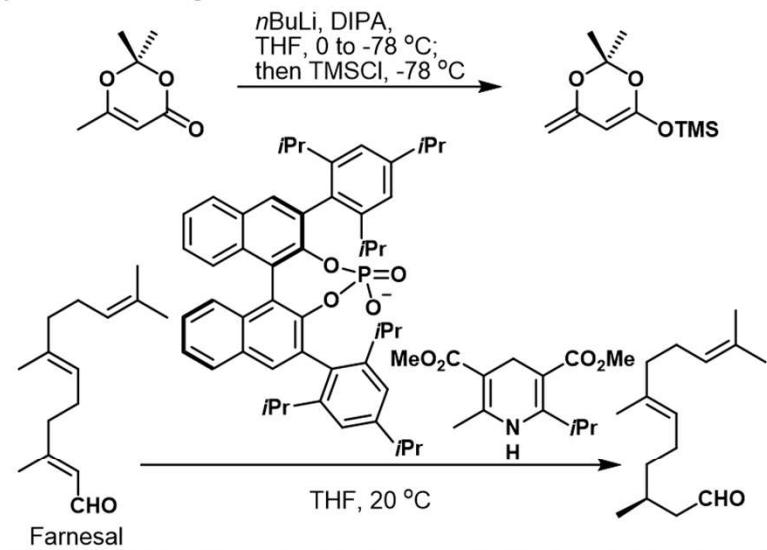
Previous syntheses:

- Grossinger (1998): 33 steps, 0.7% overall yield (racemic)
- Deslongchamps (2003): 23 steps to anhydrochatancin, 1.2% overall yield (enantioselective)
 - utilized a transannular Diels-Alder approach
 - 1.9% yield over 15 steps to TADA precursor

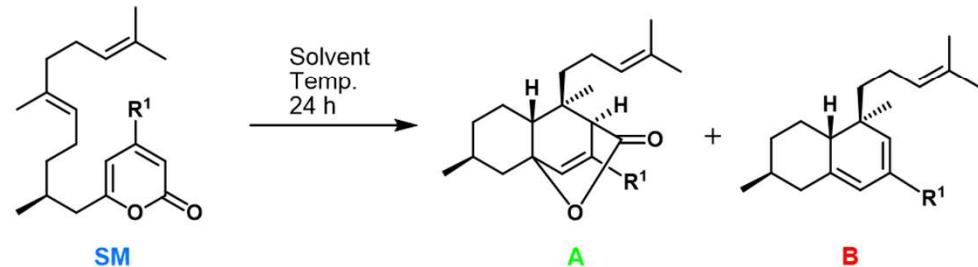


- isolated by Sato and co-worker from *Sarcophyton* species of marine corals off the coast of Okinawa, Japan
- Platelet-activating factor (PAF) is a small-molecule phospholipid mediator of numerous biological processes, including platelet aggregation, smooth muscle contraction and hypotension
 - altered levels of PAF have been implicated in numerous diseases, including respiratory and cardiovascular diseases
- inhibits PAF induced platelet aggregation ($IC_{50} = 2.2 \mu M$) and PAF receptor binding ($IC_{50} = 0.32 \mu M$) without effecting platelet aggregation induced by arachidonic acid, ADP, or collagen
- two *cis*-decalin motifs locked into place with a hemiketal bridge is crucial for biological effects

Synthesis of starting materials:



Maimone's cycloaddition optimization:



R ¹	Solvent	Temp. (°C)	Ratio (SM : A : B)
OH	toluene	165	1 : 0 : 0
OTf	toluene	100	1 : 0 : 0.15
CO ₂ Me	heptane	100	1 : 0.9 : 0.02
CO ₂ Me	PhCF ₃	100	1 : 1.4 : 0.07
CO ₂ Me	MeCN	100	1 : 2.4 : 0.4
CO ₂ Me	DMF	100	1 : 3.1 : 0.7

References:

- T. J. Maimone, *Angew. Chem. Int. Ed.* **2015**, 54, 1223
 P. Deslongchamps, *J. Org. Chem.* **2003**, 68, 6847
 E. M. Carreira, *J. Org. Chem.* **2003**, 68, 9274
 B. List, *Angew. Chem. Int. Ed.* **2006**, 45, 4193

