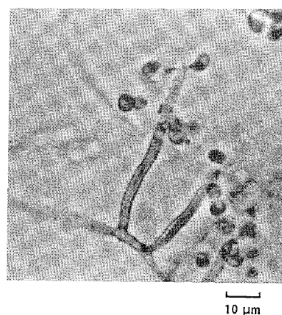


Novel diterpenoid isolated in 1987 by Ando and coworkers at Fujisawa Pharmaceutical Co. from fungal strain *Virgaria nigra*, isolated from soil collected at the foot of Mt. Aso, Japan:

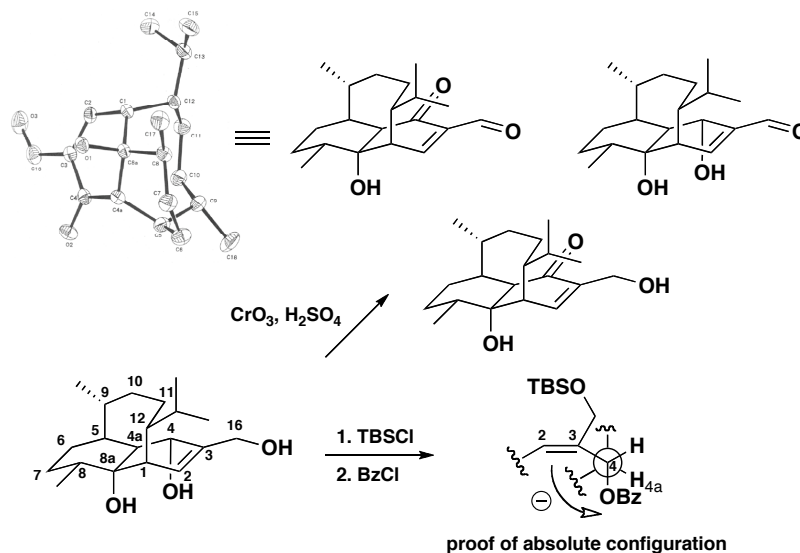


Virgaria nigra also found in US, Canada, and Cuba

Biological activity: antihypertensive, inhibition of platelet aggregation (rabbit and human), induces contraction of aortic smooth muscle (rat) through Ca^{2+} ion channel agonist activity, TNF antagonist for possible treatment of endotoxic shock, inflammation, muscle atrophy, progression of ARC to AIDS, autoimmune diseases, arthritis...

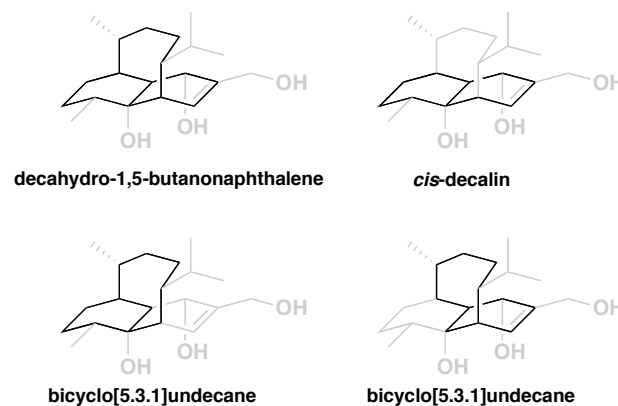
J. Antibiot. **1988**, *41*, 25 – 30 and 31 – 35.
Org. Prep. Proc. Int., **2007**, *39*, 311 – 353.

Publication breakdown: 16 studies toward, 4 dissertations, 8 patents, 1 review, 0 total syntheses.



J. Org. Chem. **1987**, *52*, 5292 – 5293.

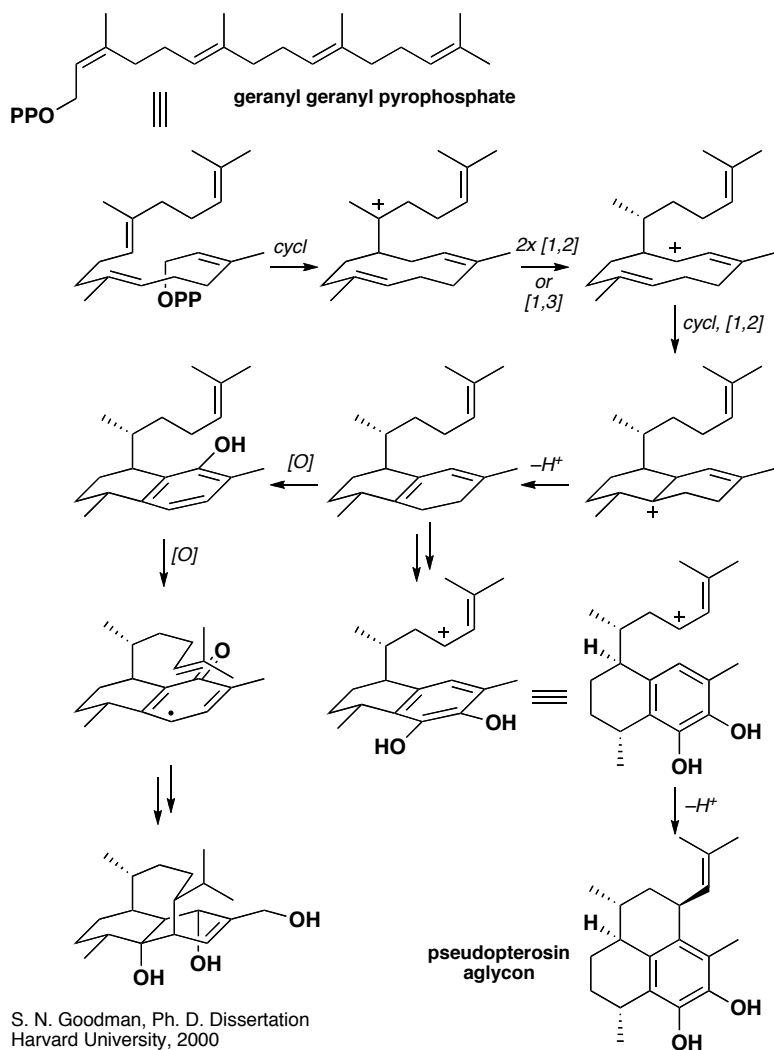
embedded core structures:



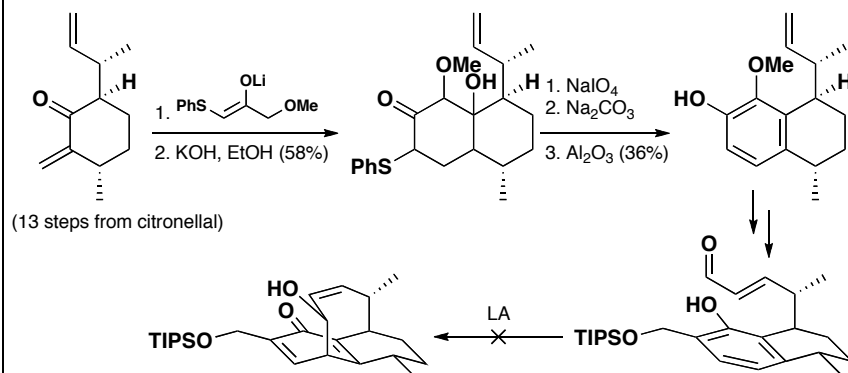
N. Z. Burns

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Proposed biosynthesis by Corey and Goodman:

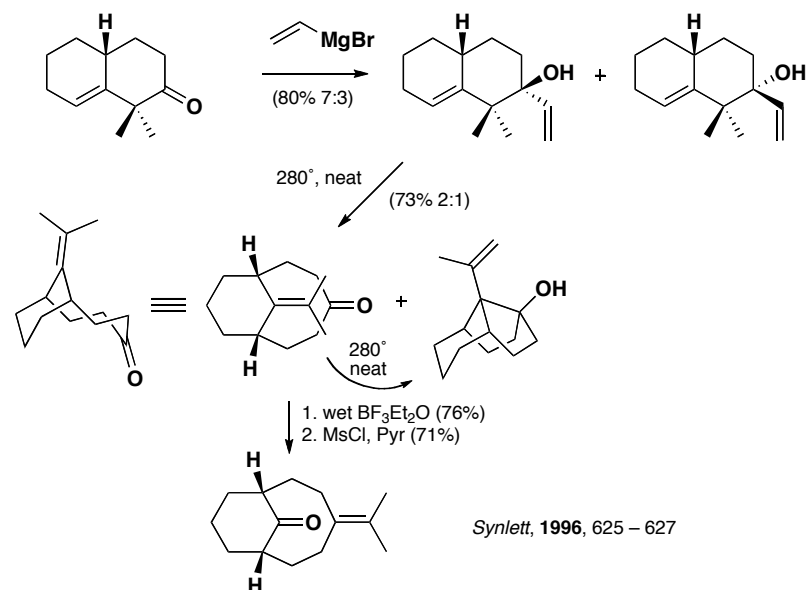


Corey biomimetic approach:

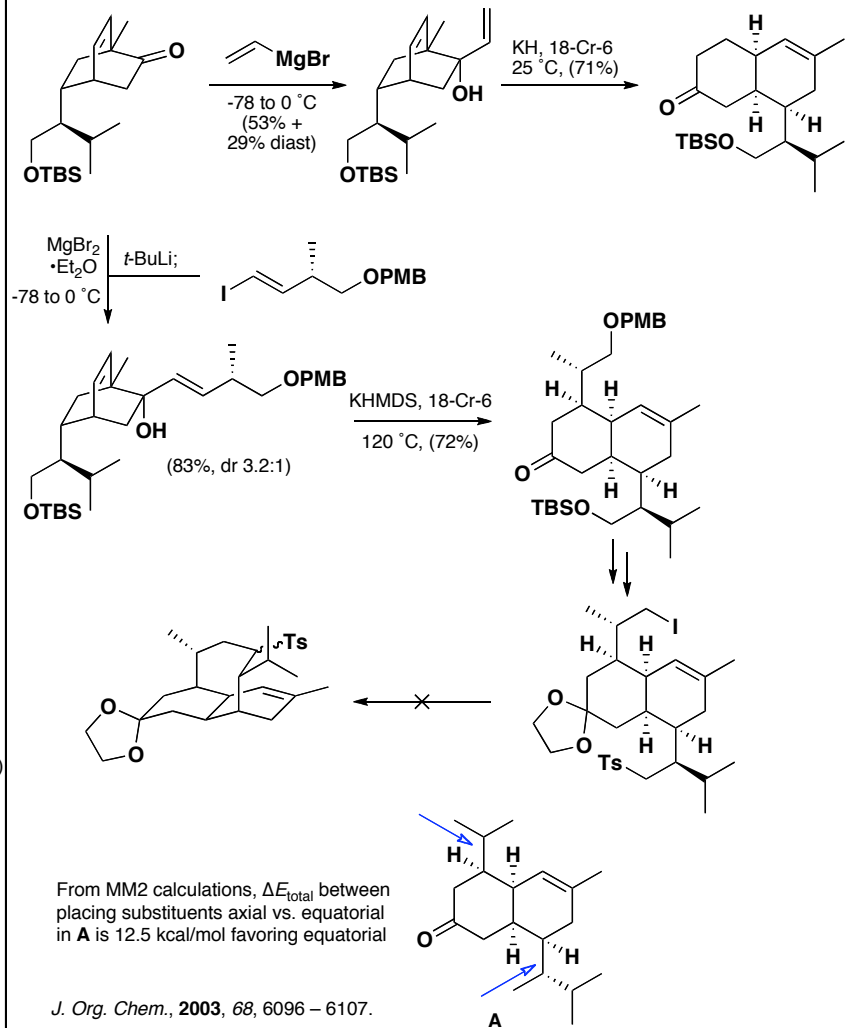
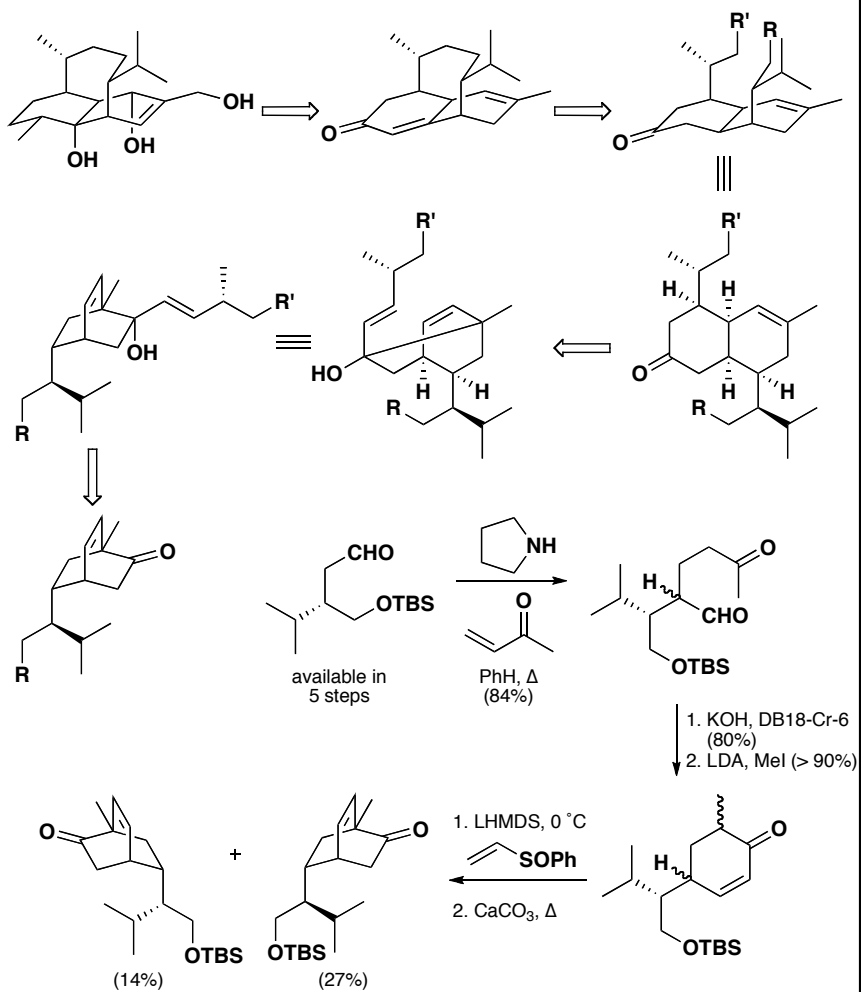


A. Palani, Harvard University, Unpublished results in:
S. N. Goodman, Ph. D. Dissertation, Harvard University, 2000

Mehta's bicyclo[5.3.1]undecane:



Paquette first generation:

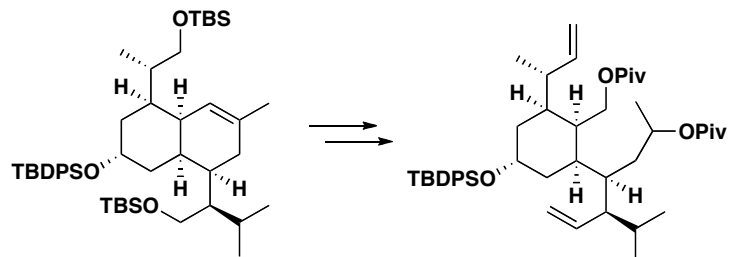
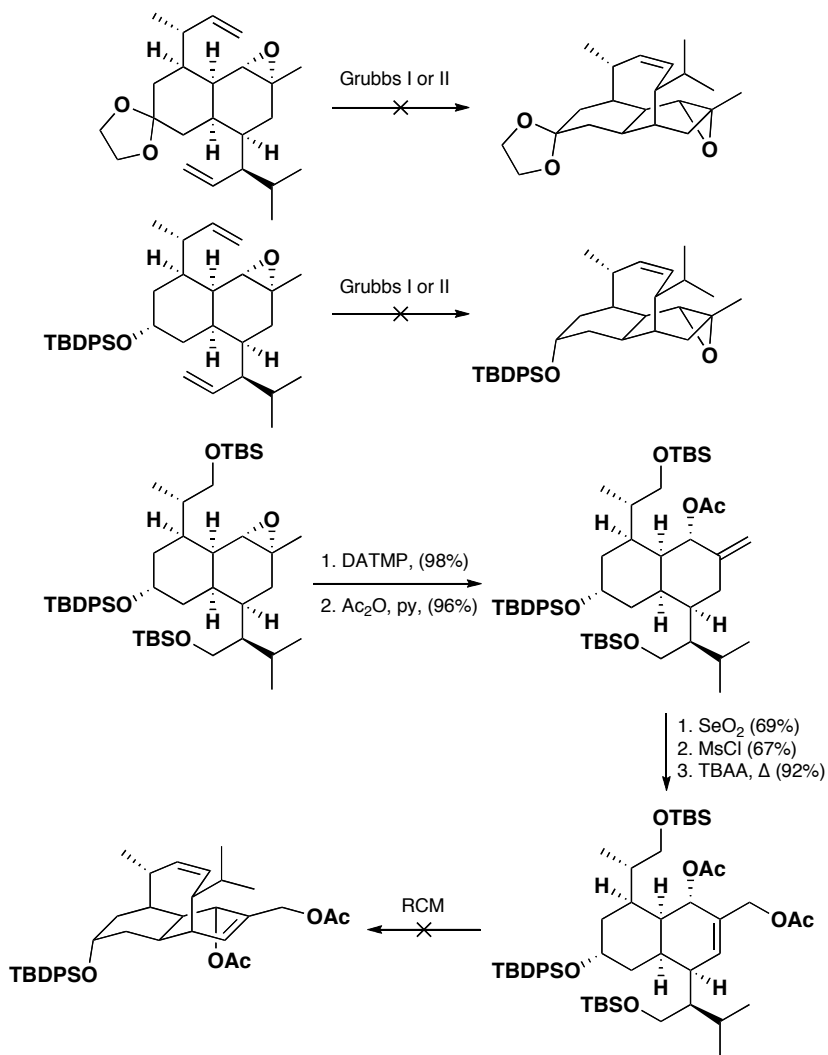


From MM2 calculations, ΔE_{total} between placing substituents axial vs. equatorial in **A** is 12.5 kcal/mol favoring equatorial

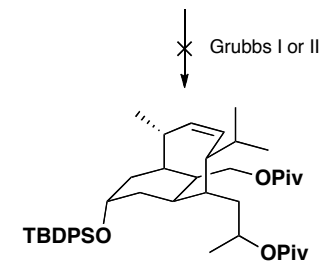
J. Org. Chem., 2003, 68, 6096 – 6107.

Vinigrol

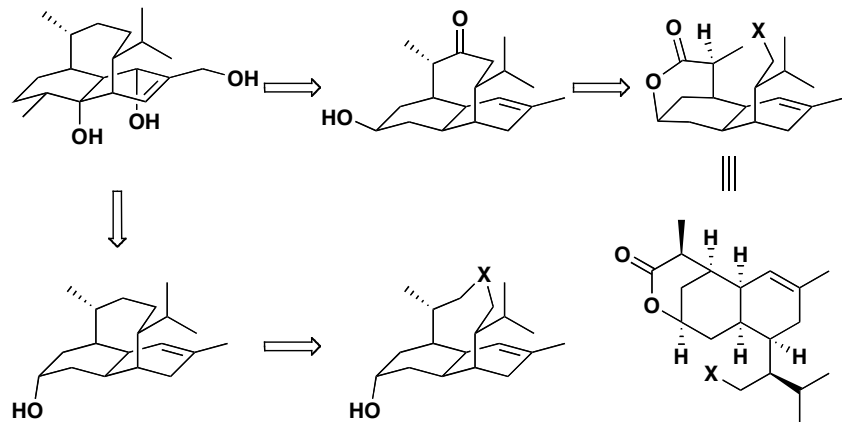
Paquette part deux:



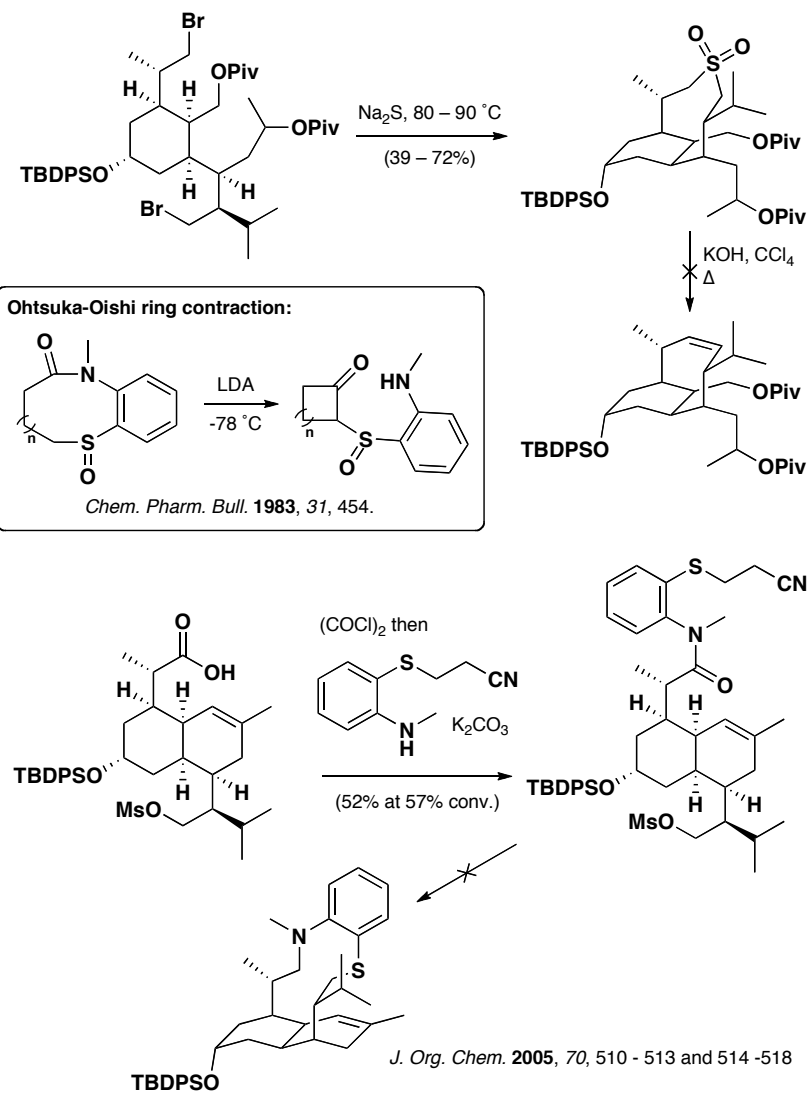
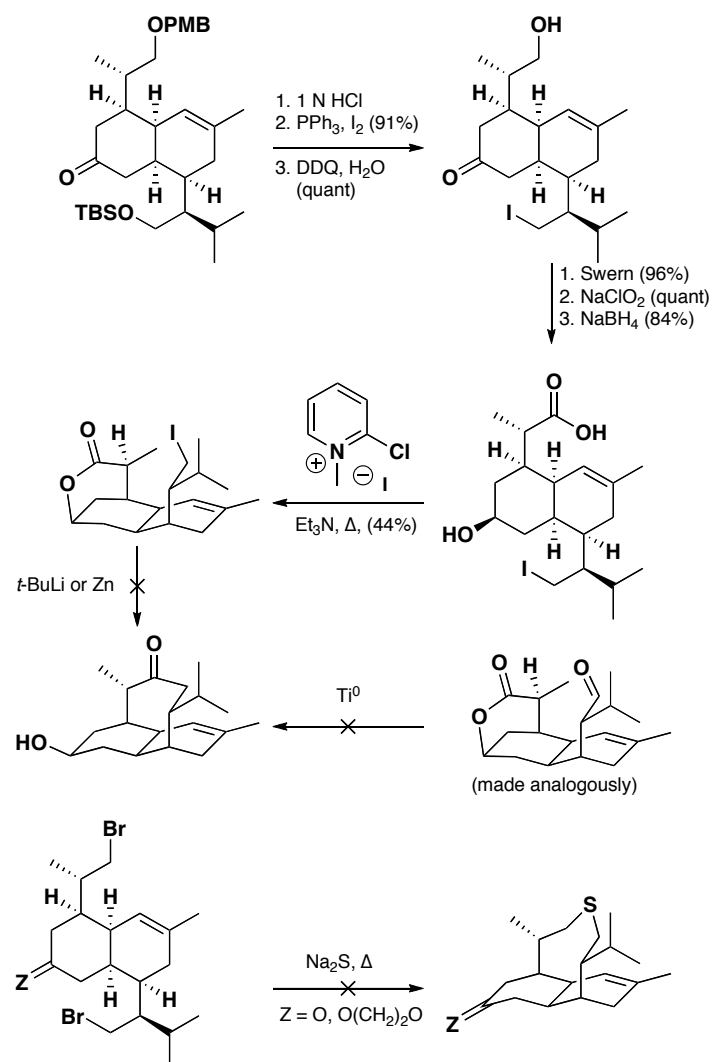
J. Org. Chem. 2005, 70, 505-509.



Paquette generation two and three:



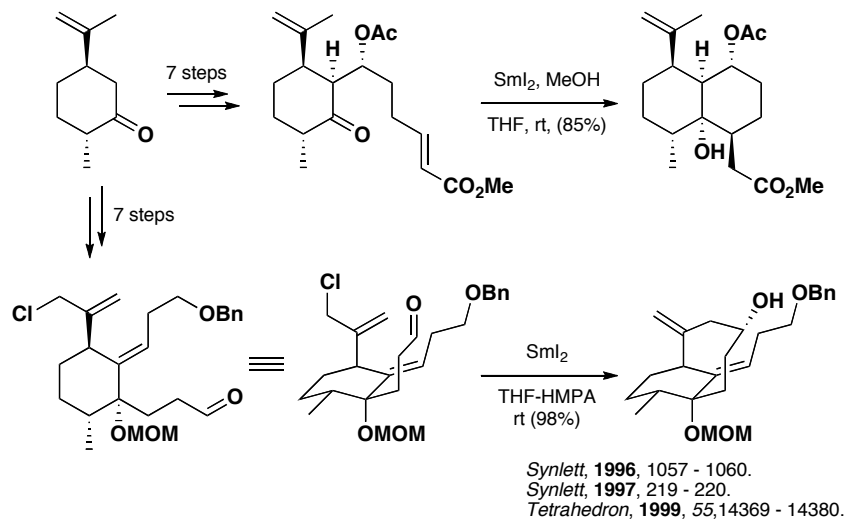
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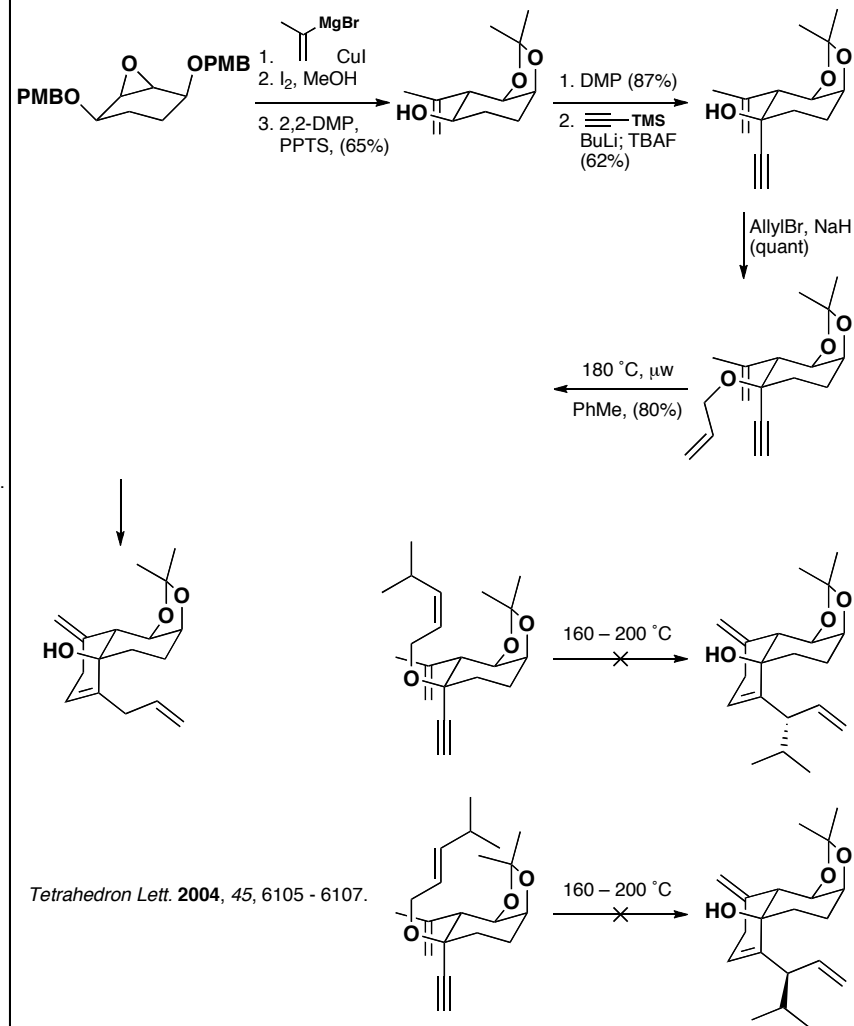
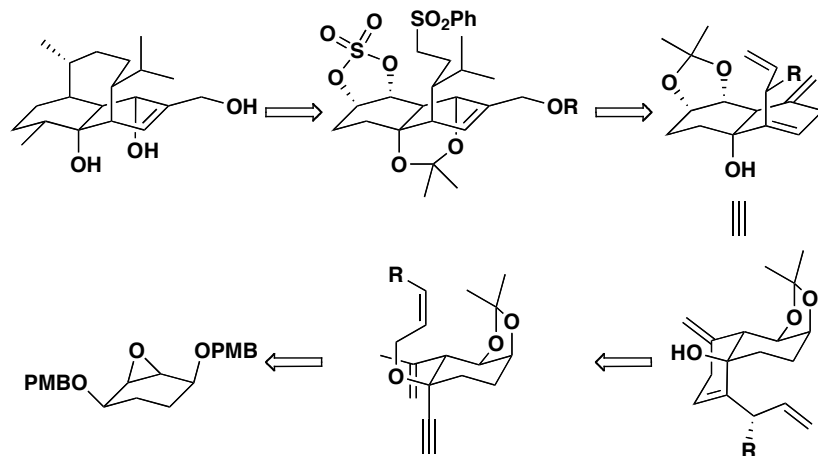
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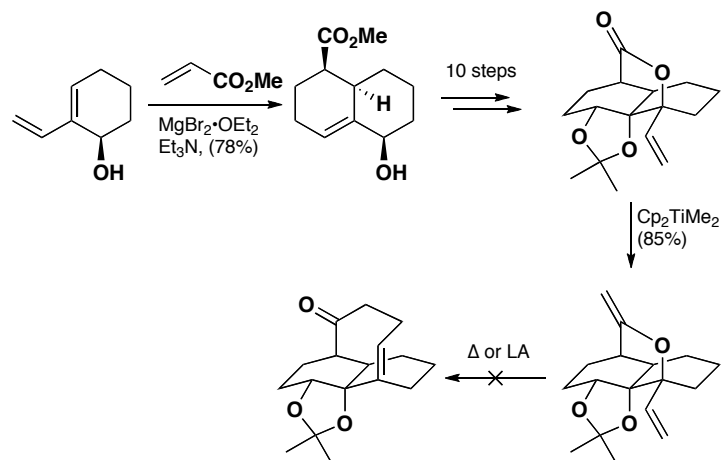
Matsuda's SMI₂ approaches:



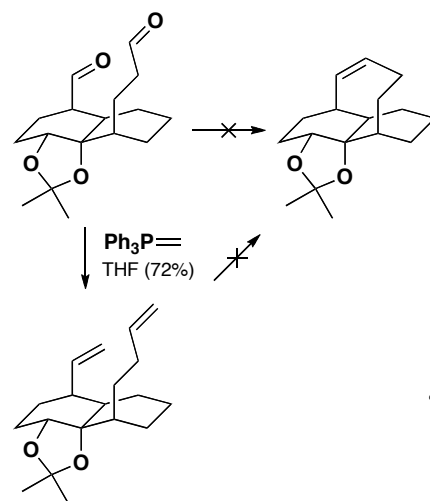
Barriault cascade:



Barriault second generation:

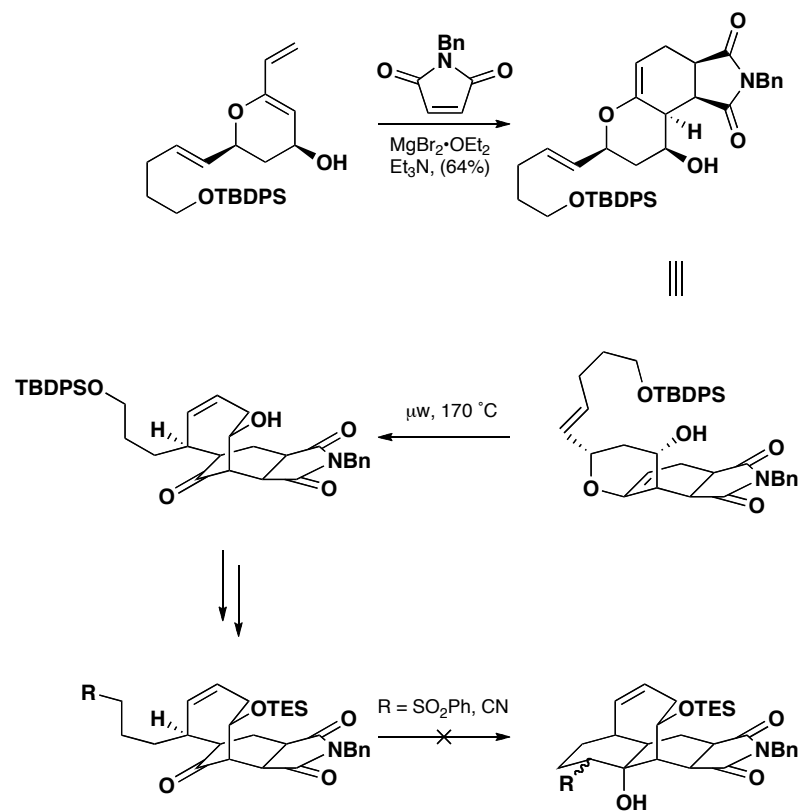


Barriault macrocyclization:



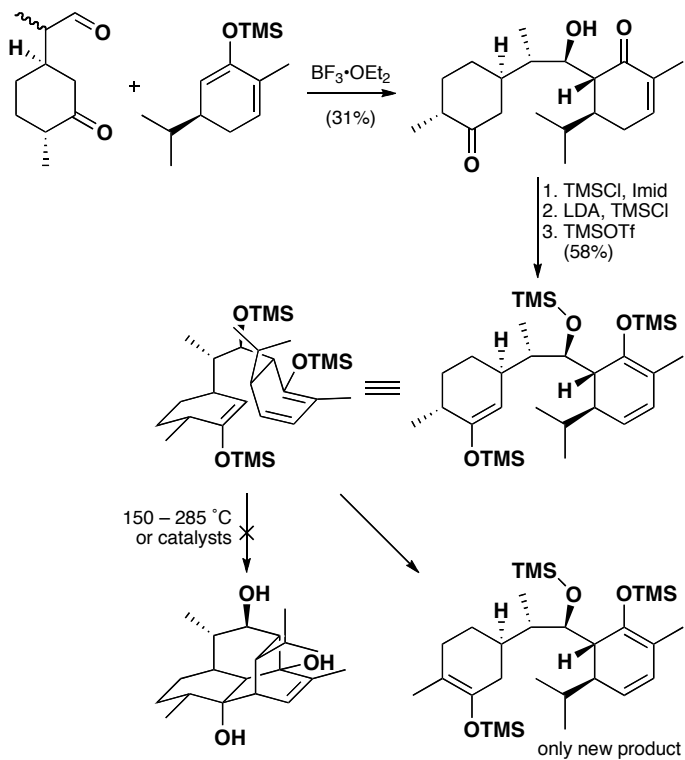
J. Org. Chem. **2005**, *70*, 8841 - 8853.

Barriault fourth generation:

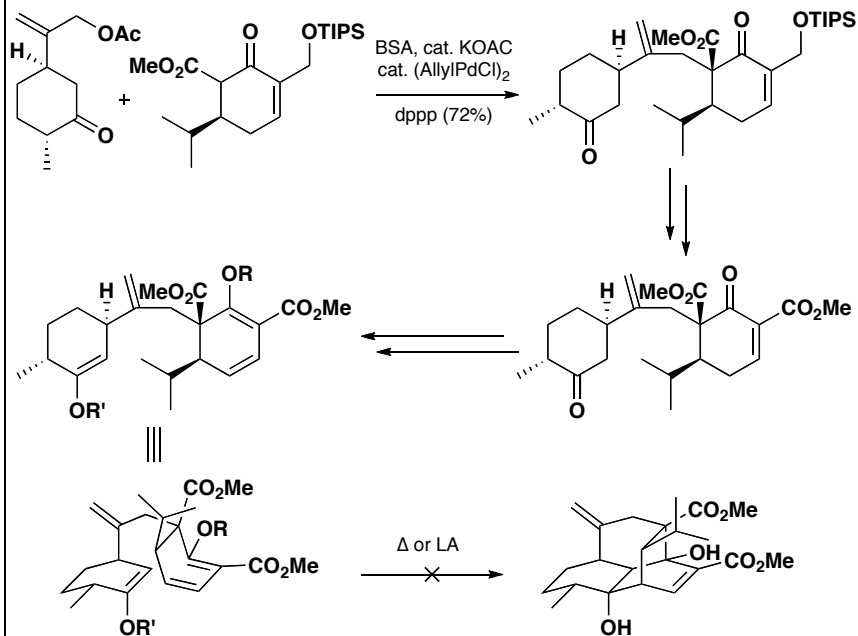
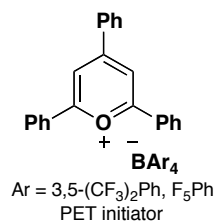
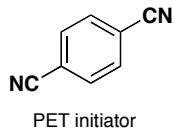
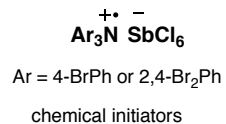


J. Org. Chem. **2005**, *70*, 8841 - 8853.

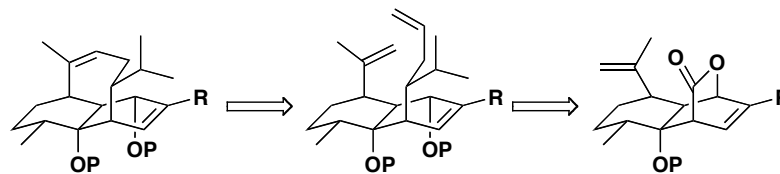
Corey IMDA approaches:



CET or PET catalysts tried for DA:



Another Corey strategy:

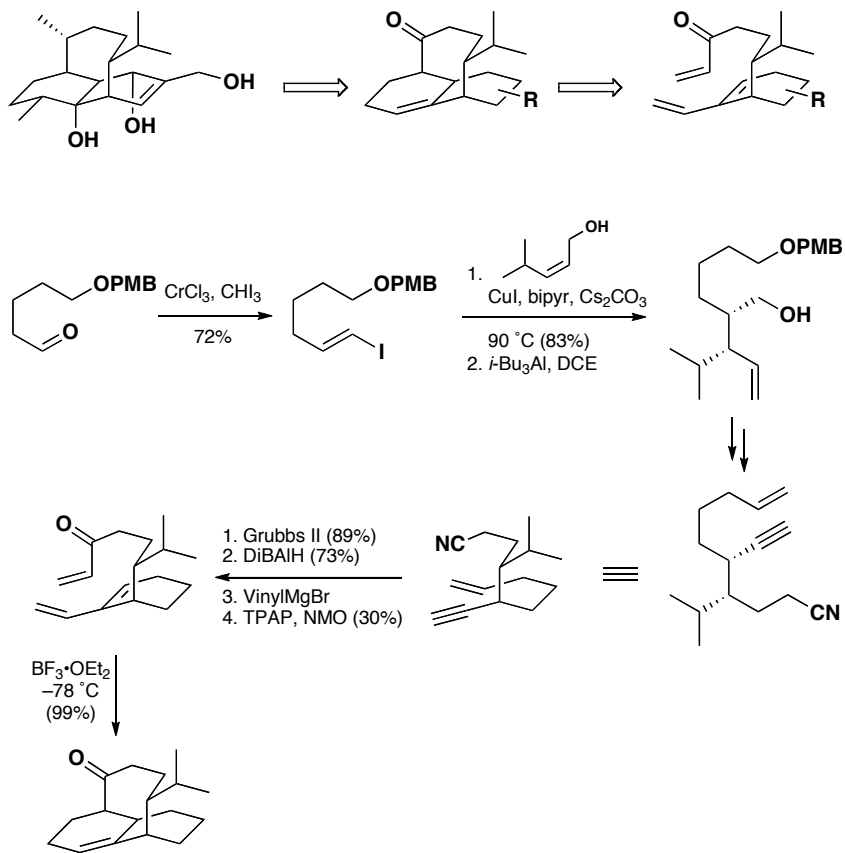


S. N. Goodman, Ph. D. Dissertation
Harvard University, 2000

N. Z. Burns

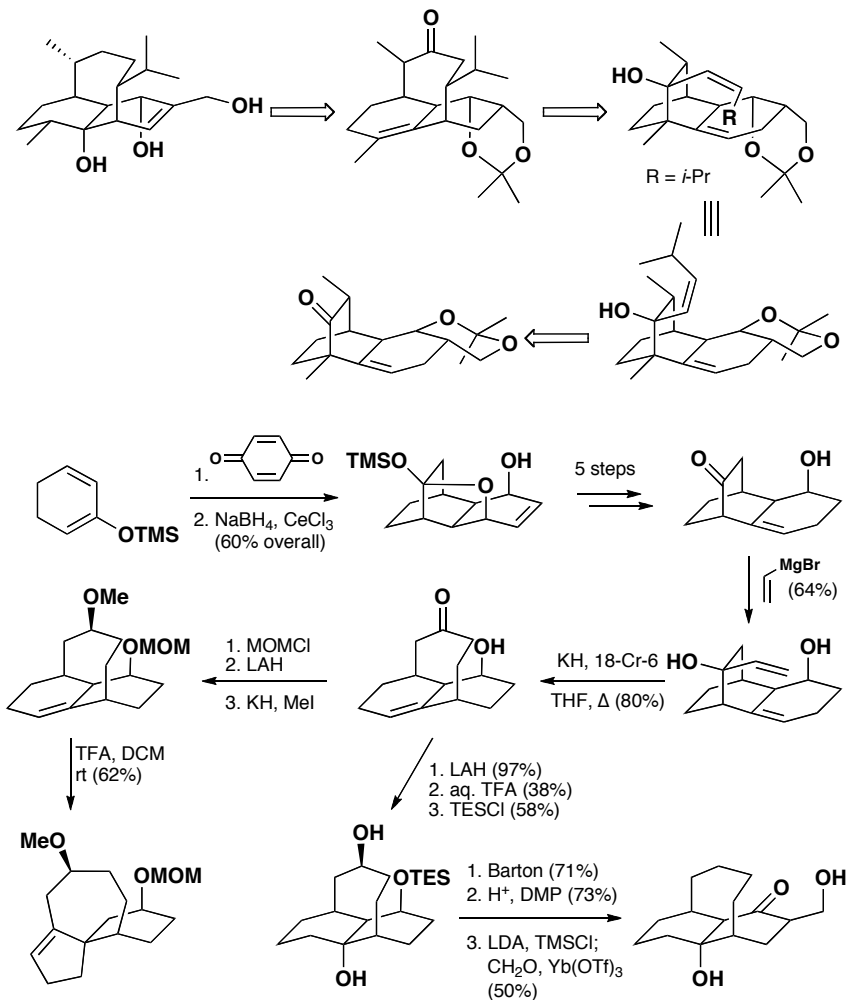
Vinigrol

Barriault assault:



Org. Lett. 2007, 9, 1545 - 1548.

Hanna's advanced intermediate:

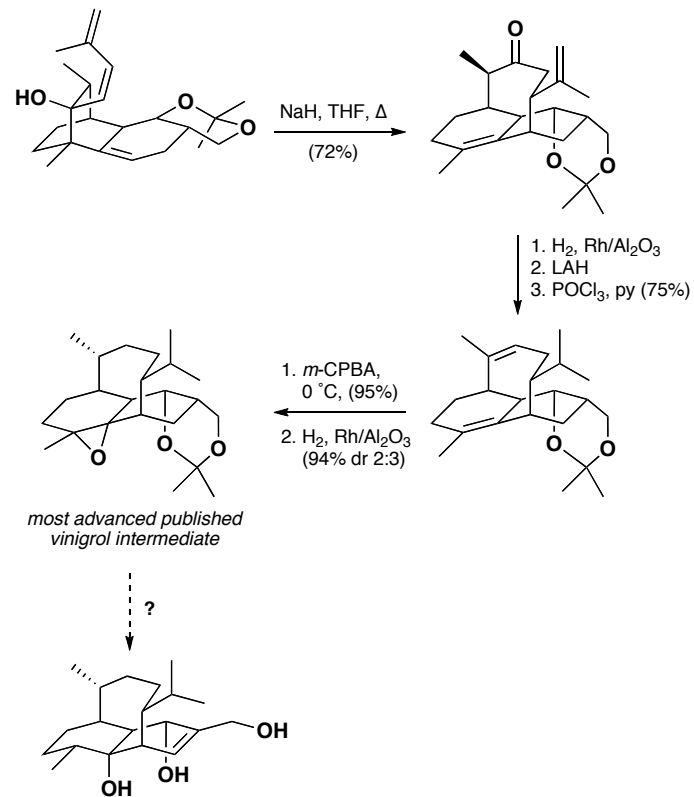
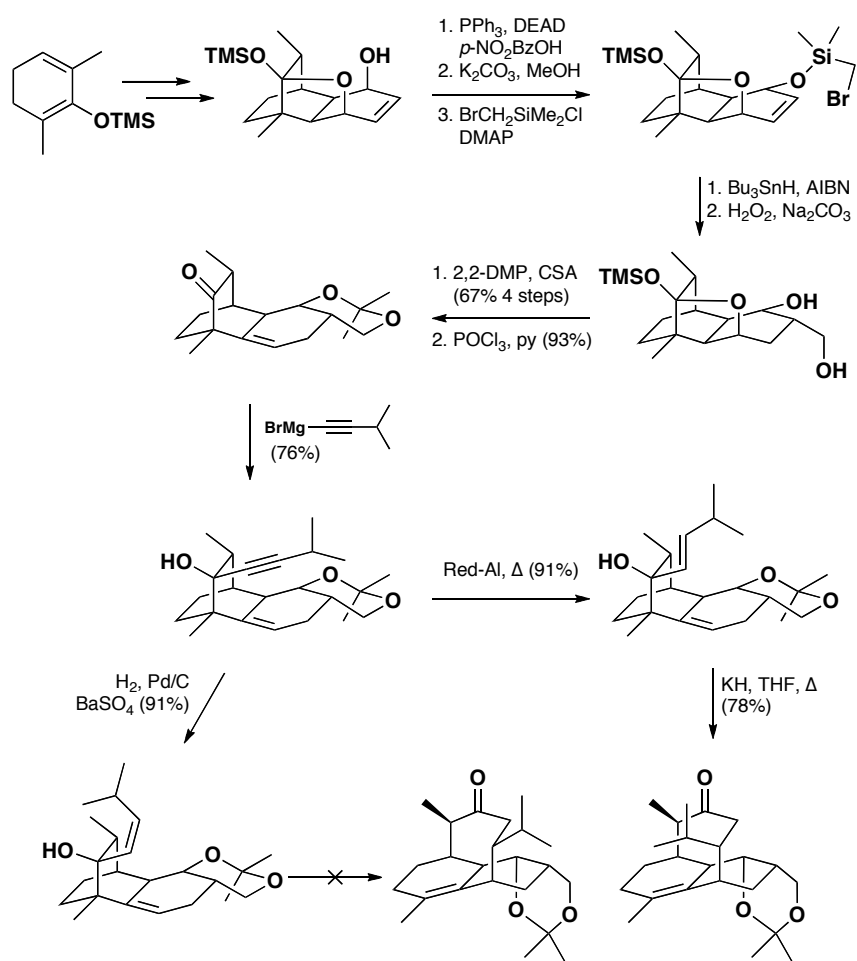


J. Org. Chem. 1993, 58, 2349 - 2350.
 J. Chem. Res. (Syn), 1996, 32 -33.

J. Org. Chem. 1997, 62, 5062 - 5068.

N. Z. Burns

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Org. Lett. 2003, 5, 1139 - 1142.