

Asymmetric Total Synthesis of (–)- Scabronine G via Intramolecular Double Michael Reaction and Prins Cyclization

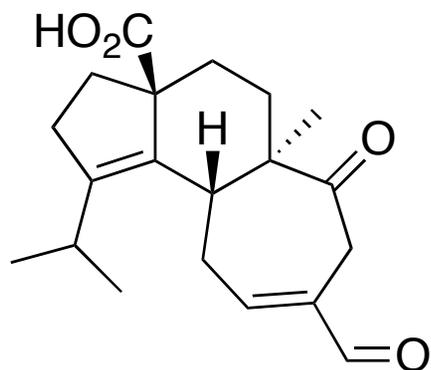
Naoki Kanoh, Kohei Sakanishi, Emiko Imori, Ken'ichi
Nishimura, Yoshiharu Iwabuchi

Org. Lett. **2011**, ASAP,
doi: 10.1021/ol200873y

Current Literature

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(-)-Scabronine G

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<http://www.mushroomexpert.com/sarcodo>

Sarcodon scabrosus is a mushroom belonging to the family *Thelephoraceae* and has a strongly bitter *taste*.

It is a widely distributed species in Europe and in North America.

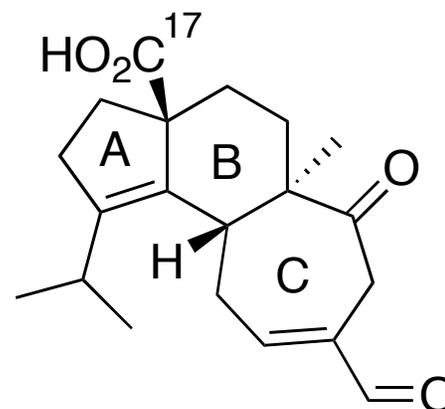
Scabronine G was isolated by Ohta and co-workers in 2001 and is shown to enhance the secretion of neurotrophic factors from 1321N1 human astrocytoma cells.

Scabronine G and their methyl ester are useful drugs to clarify the mechanisms underlying the synthesis and secretion of neurotrophic factors.

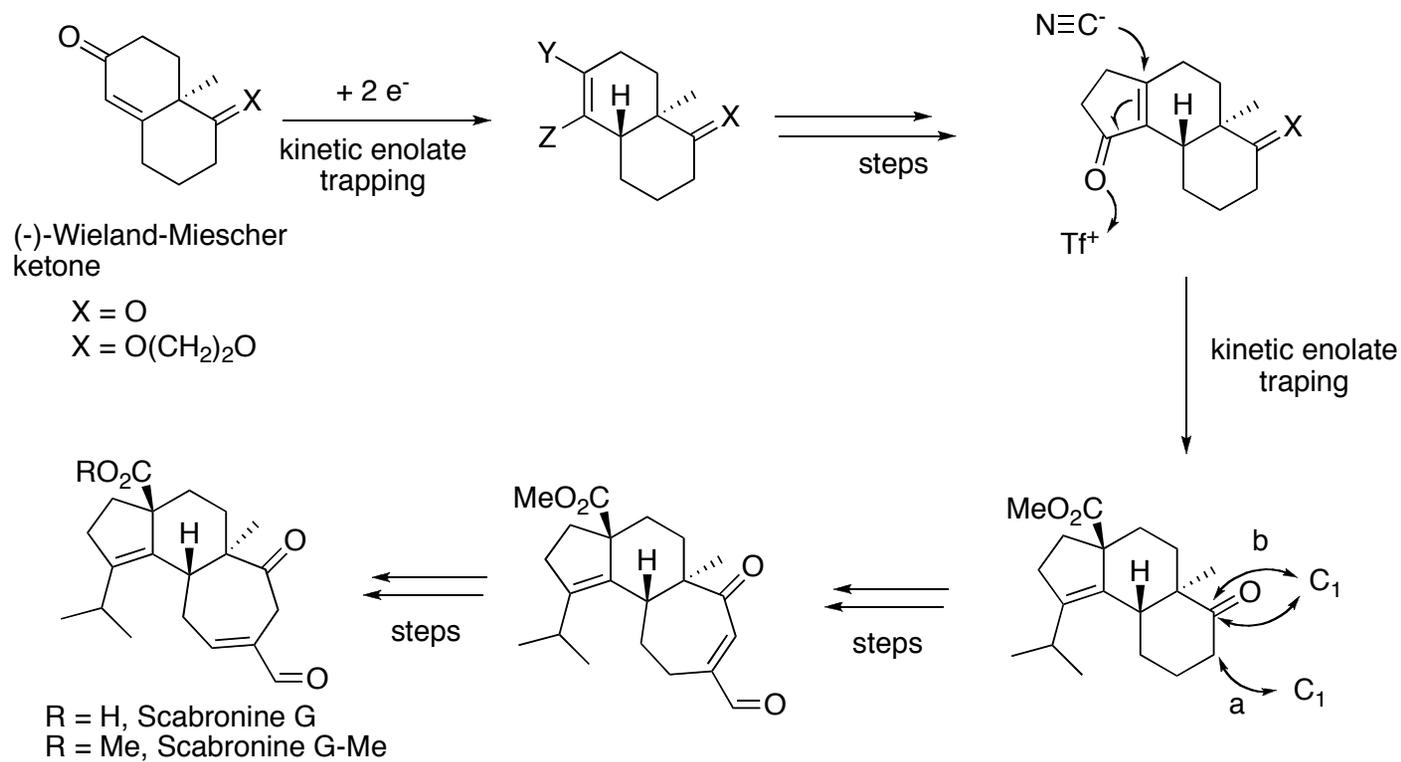
Y. Obara and co-workers *Y. Mol. Pharmacol.* **2001**, 59, 1287.

(-)-Scabronine G

- Cyathane diterpene natural product
- tricyclic 5-6-7 ring system
- trans- fused 6-7 ring
- angular C-17 carboxyl group

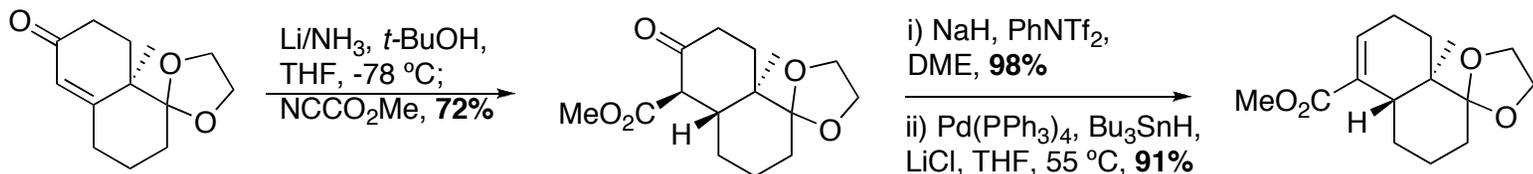


Previous total synthesis (-)-Scabronine G

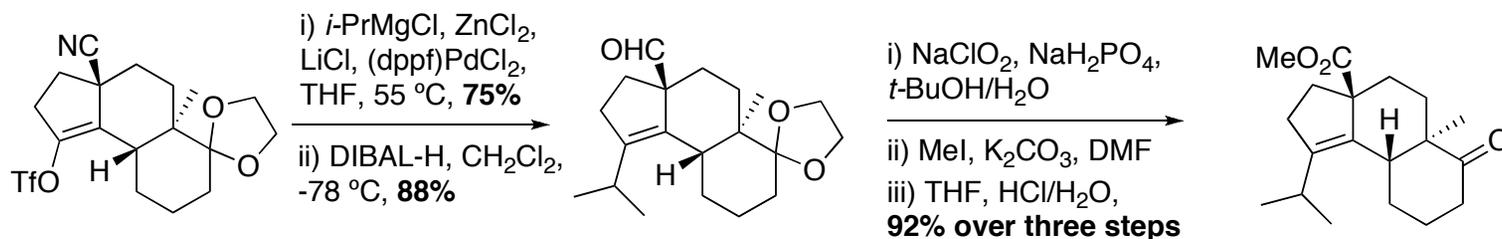
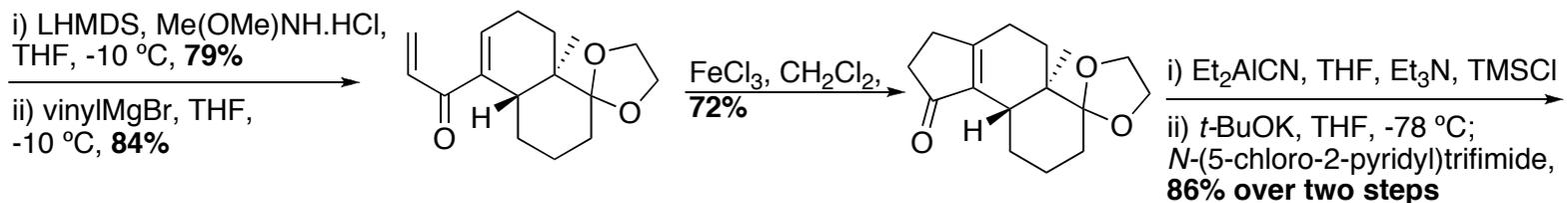


J. Danishefsky and co-workers J. Am. Chem. Soc. **2005**, *127*, 13514

Previous total synthesis (-)-Scabronine G

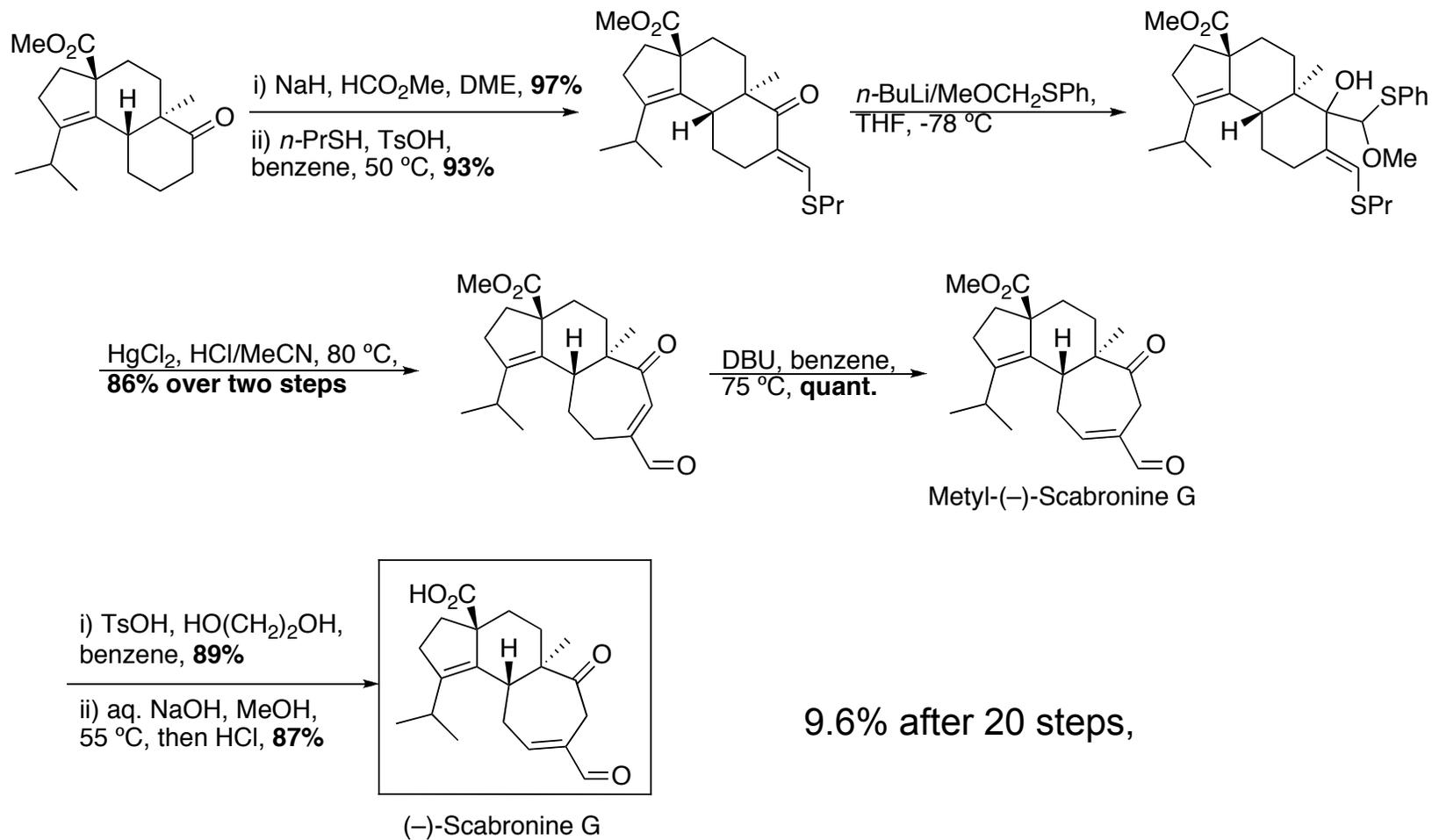


(-)-Wieland-Miescher ketone



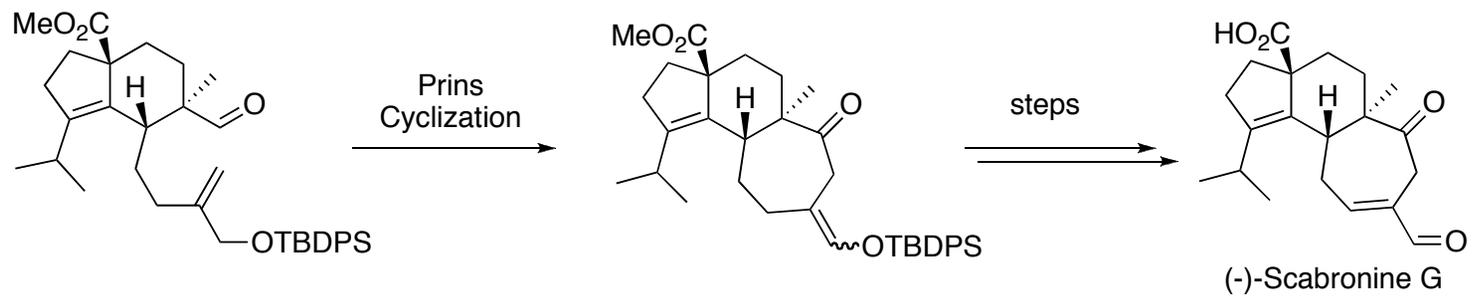
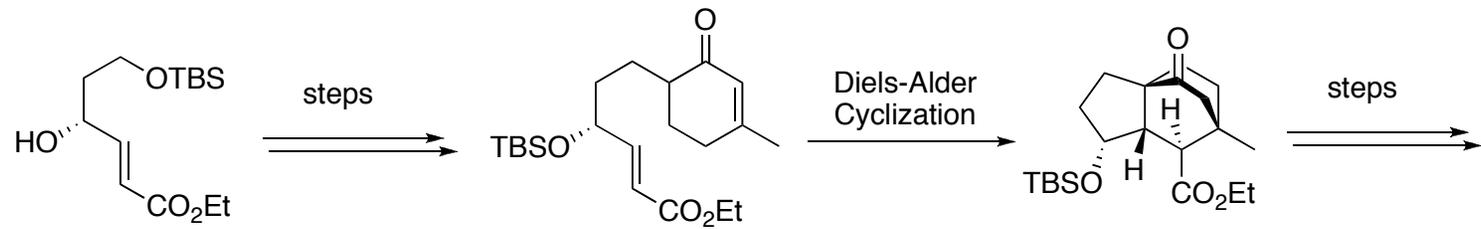
J. Danishefsky and co-workers *J. Am. Chem. Soc.* **2005**, *127*, 13514

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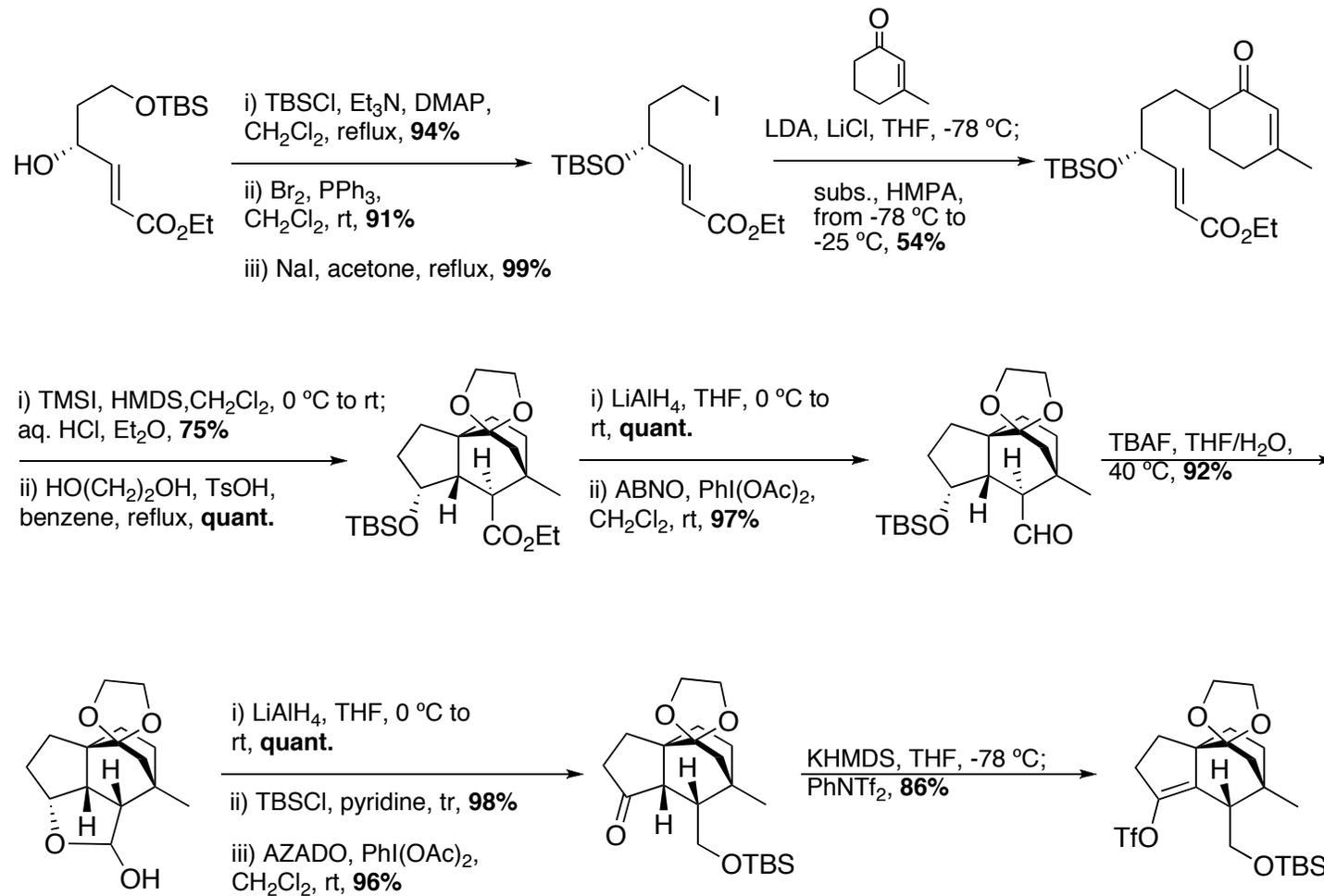
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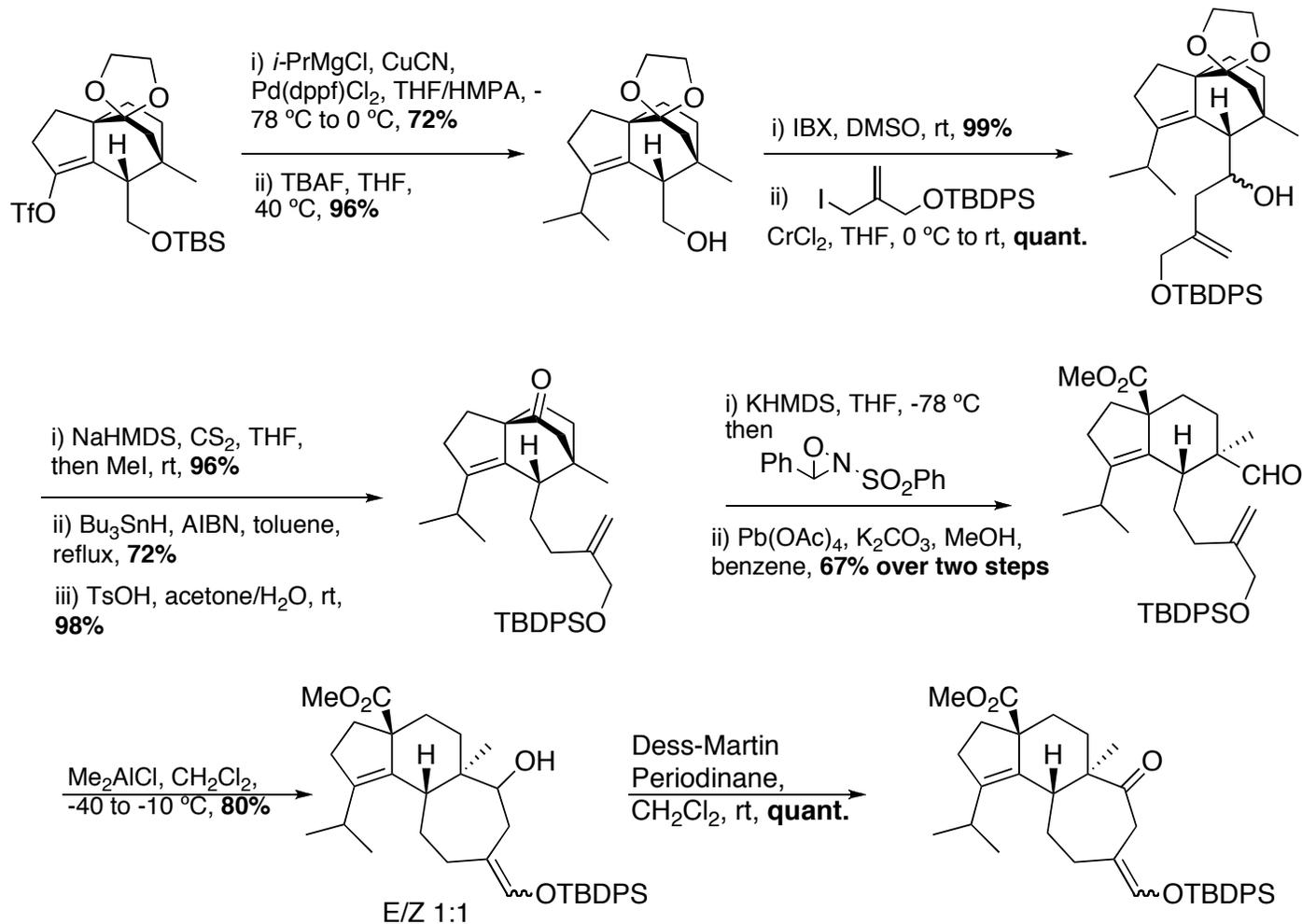
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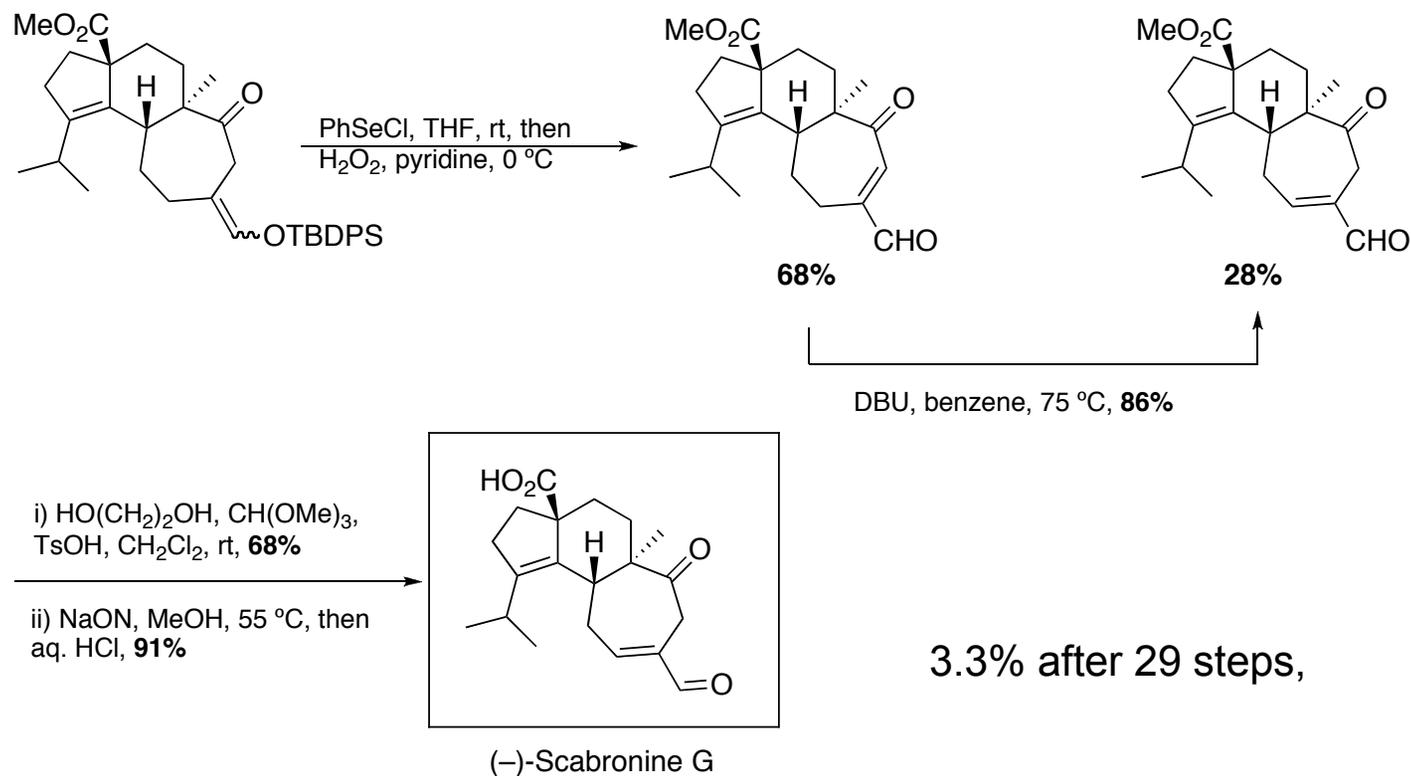
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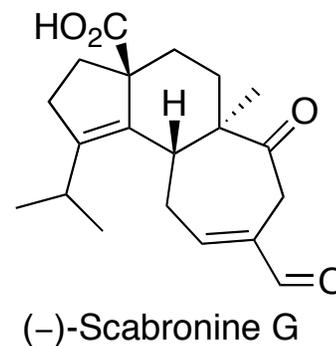
N. Kanoh and co-workers *Org. Lett.* **2011**, ASAP

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N. Kanoh and co-workers *Org. Lett.* **2011**, ASAP

Summary



- (-)-Scabronine G was synthesized in 29 steps in 3.2%
- Key transformations include selective Diels-Alder cyclization and Prins cyclization