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The Total Synthesis of Racemic Talatisamine

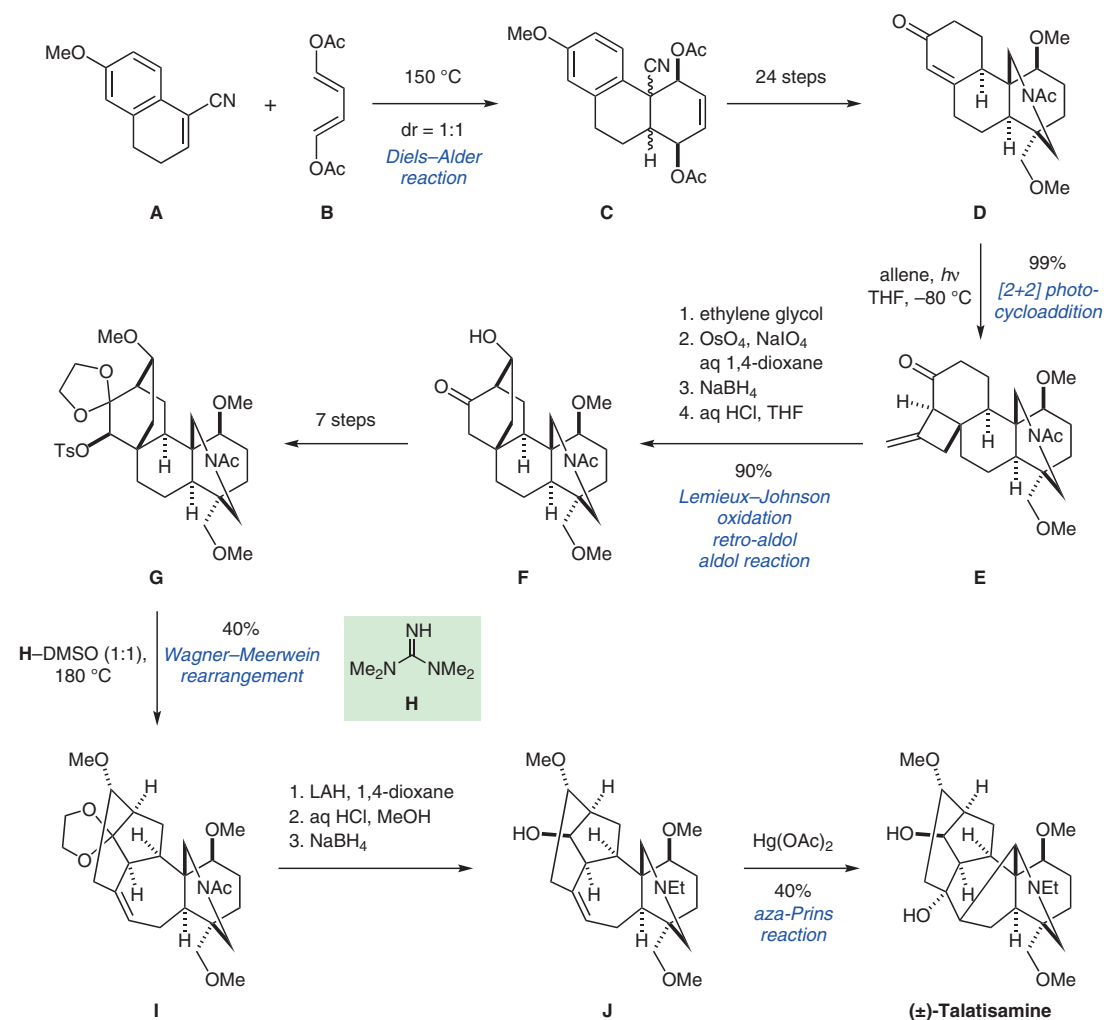
Pure Appl. Chem. **1975**, *41*, 93–112, DOI: 10.1351/pac197541010093.

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Total Synthesis of Talatisamine, a Delphinine Type Alkaloid

J. Am. Chem. Soc. **1974**, *96*, 4990–4992, DOI: 10.1021/ja00822a048.

Total Synthesis of (±)-Talatisamine



Significance: (–)-Talatisamine is a potassium ion channel inhibitor isolated from an *aconitum* species. Wiesner and co-workers reported the first total synthesis of (±)-talatisamine. Its preparation was facilitated by synthetic studies on degradation products of natural (–)-talatisamine. Other diterpene alkaloids prepared by the Wiesner group include atisine and napelline.

Comment: The tricyclic core of (±)-talatisamine is assembled by a Diels–Alder reaction. Ring expansion from the cyclobutane in **E** via retro-aldol–aldol reaction furnishes the bicyclo[2.2.2]octane motif within **F**. The atisine-type scaffold **G** is converted into the delphinine-type skeleton **I** by Wagner–Meerwein rearrangement in analogy to Wiesner’s biosynthetic proposal. An oxidation followed by aza-Prins reaction furnishes the hexacyclic 6/6/5/6/5/6-membered-ring scaffold of (±)-talatisamine.

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Synthesis of Natural Products

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(±)-talatisamine

Diels–Alder reaction

[2+2] photocycloaddition

Lemieux–Johnson oxidation

retro-aldol–aldol reaction

Wagner–Meerwein rearrangement

aza-Prins reaction

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