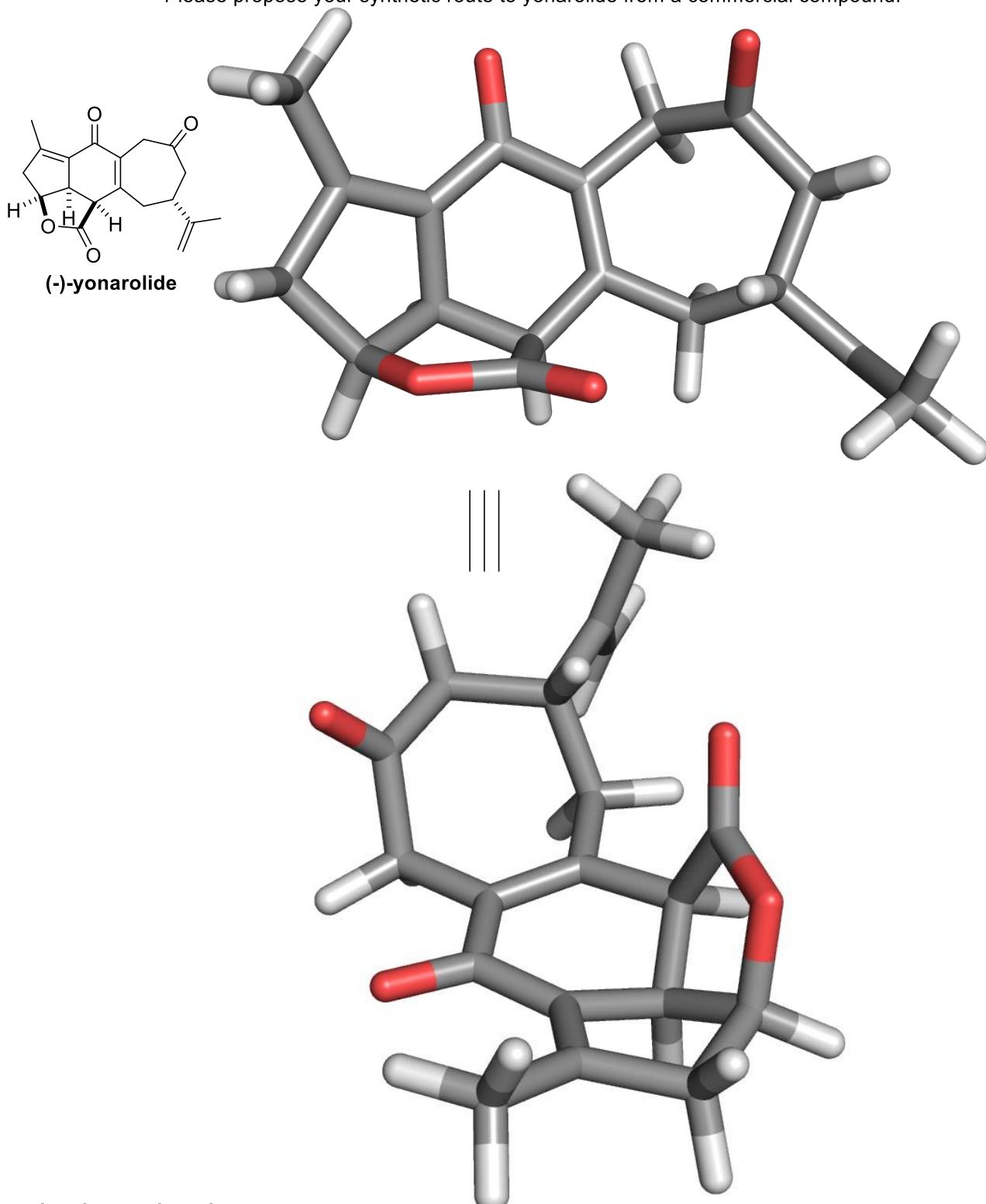


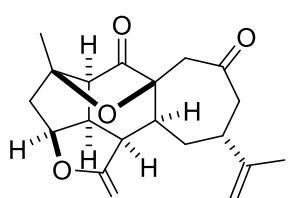
Problem Session (13)

2021/09/18 MASANORI NAGATOMO

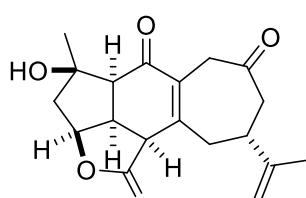
Please propose your synthetic route to yonarolide from a commercial compound.



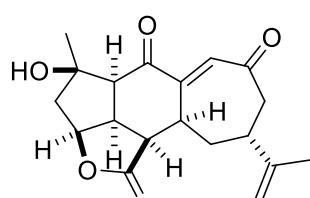
related natural products



sinulochmodin C



scabrolide A



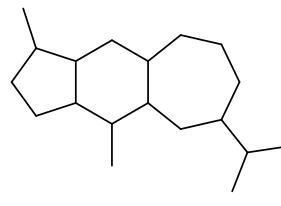
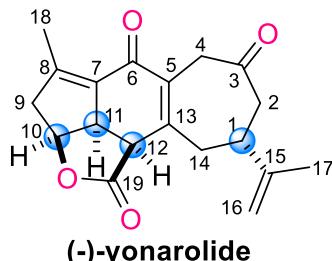
scabrolide B

Problem Session (13)- Answer

2021/09/18 MASANORI NAGATOMO

1

Synthetic Plan of (-)-yonarolide



yonarane skeleton

• **Isolation:** Isolated from Okinawan soft coral of the genus *Sinularia* along with 5-*epi*-sinuleptolide and 5,11-*diepi*-sinuleptolide. Iguchi, K.; Kajiyama, K.; Yamada, Y. *Tetrahedron Lett.* **1995**, 36, 8807.

• **Bioactivities:** Yonarolide have a potential as an anti-inflammatory agent.

• **Structural features:** Norditerpenoid having a unique tricyclo[7.5.0.0^{3,7}]tetradecane chemical structure called the yonarane skeleton.

The absolute stereochemistry of scabrolide A was determined by comparison of the calculated and measured ECD values and by total synthesis, but that of yonarolide has not yet been directly determined.

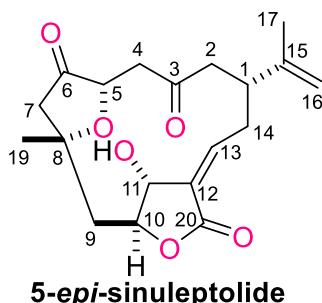
Total synthesis

No report in peer-reviewed articles.

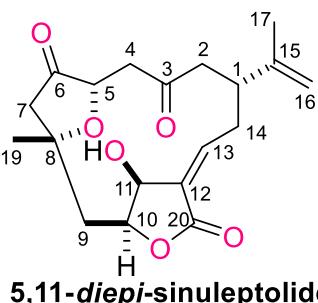
The total synthesis of the related natural product scabrolide A has been reported by the Stoltz group:

Hafeman, N. J.; Loskot, S. A.; Reimann, C. E.; Pritchett, B. P.; Virgil, S. C.; Stoltz, B. M. *J. Am. Chem. Soc.* **2020**, 142, 8585.

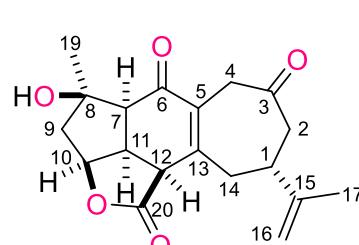
Related natural products



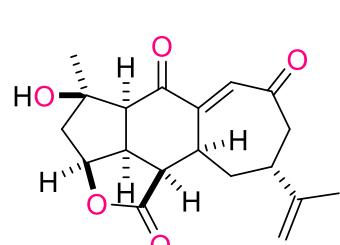
5-*epi*-sinuleptolide



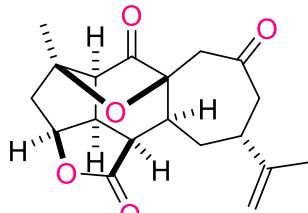
5,11-*diepi*-sinuleptolide



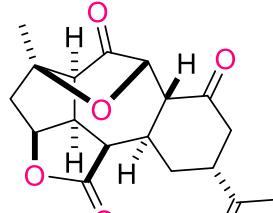
scabrolide A



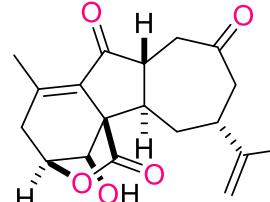
scabrolide B



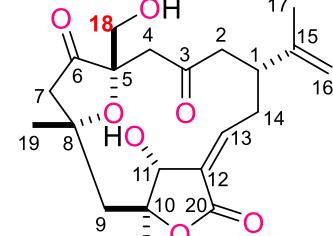
sinulochmodin C



ineleganolide

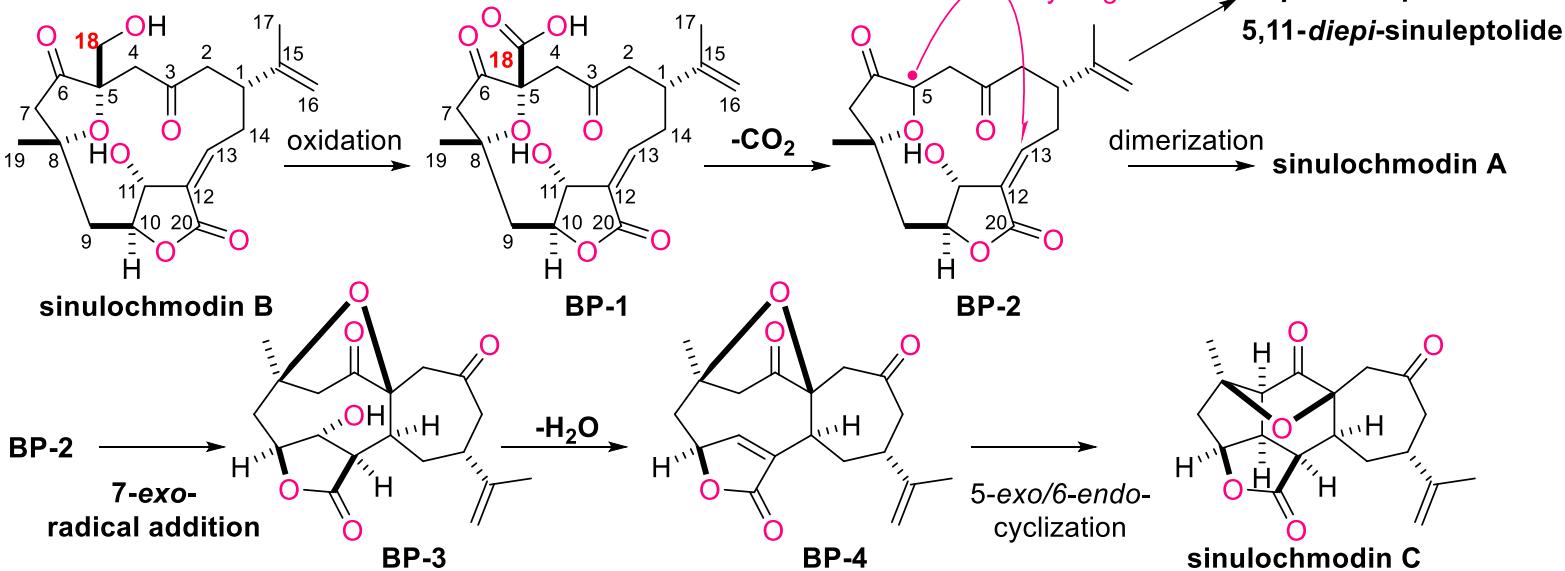


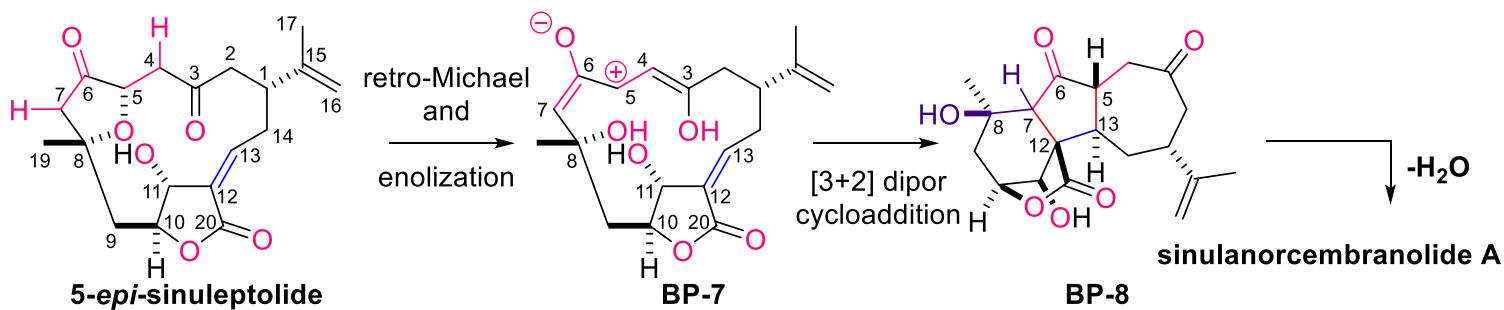
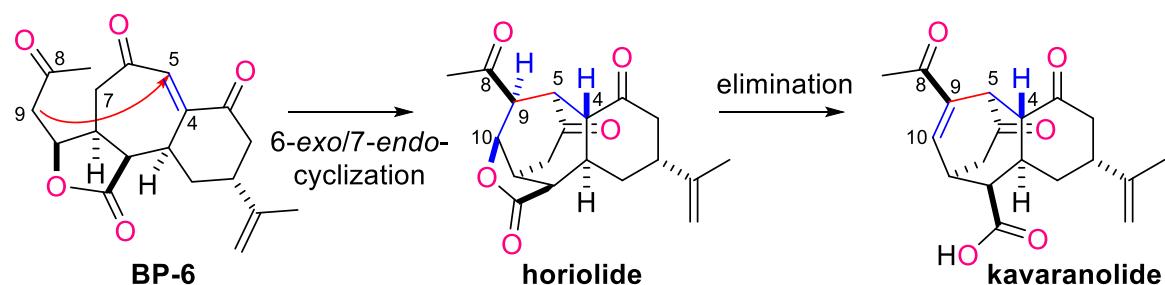
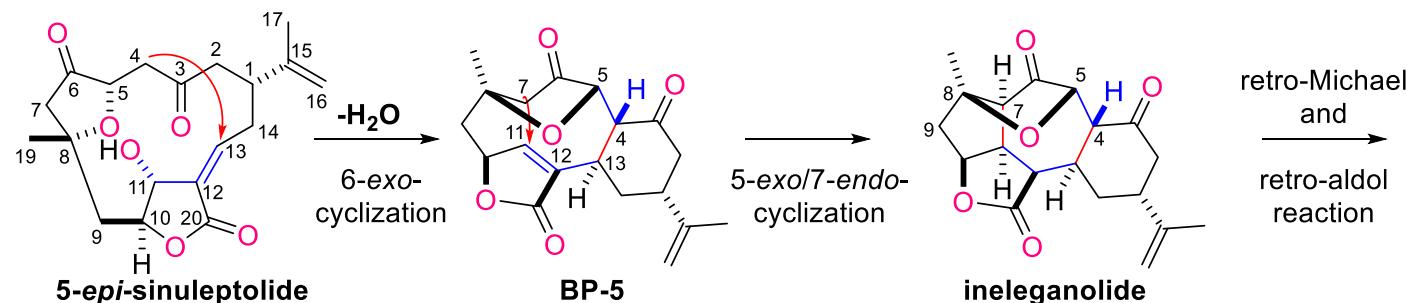
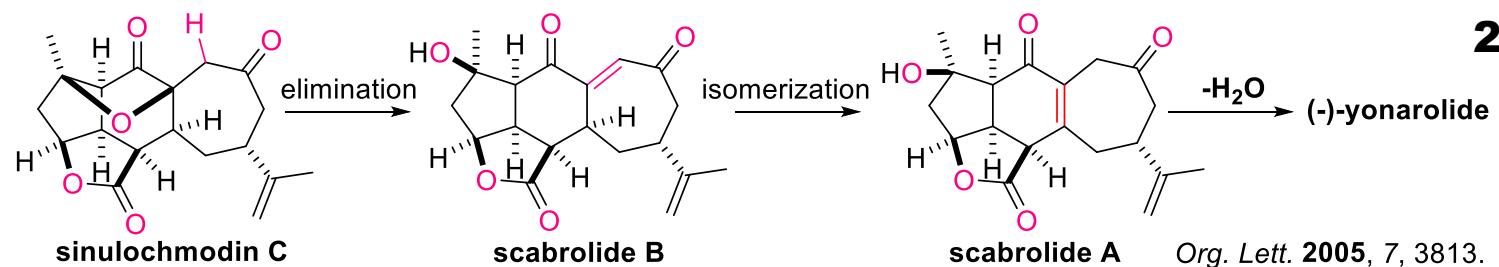
sinulanorcembranolide



sinulochmodin B

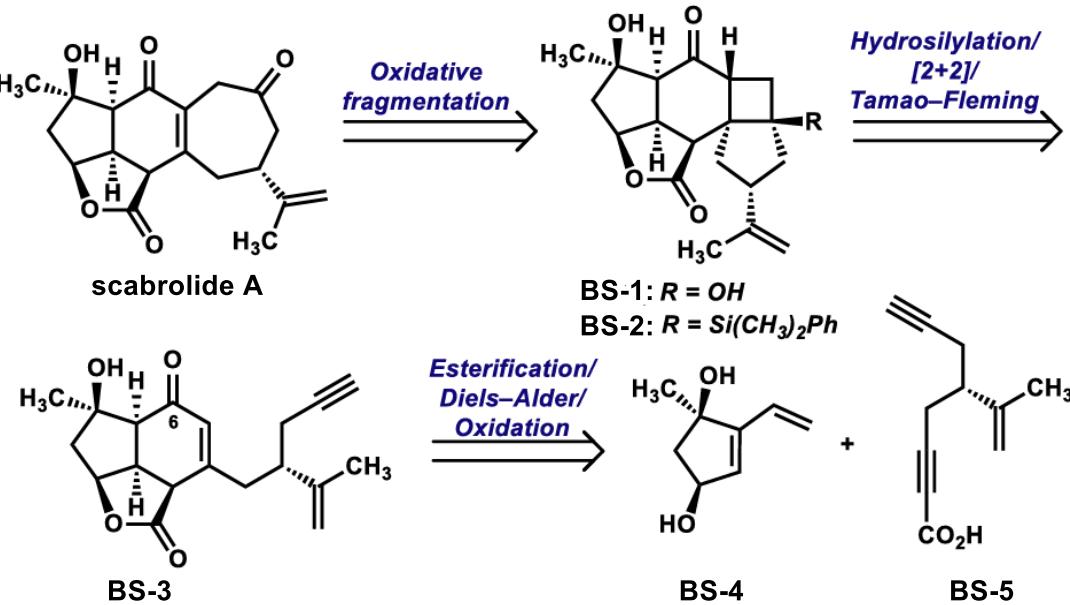
Proposed biosynthesis





Chem. Rev. 2017, 117, 7878.

• The 21-step total synthesis of scabrolide A by Stoltz group: See, 200606_PS_Takahiro_Watanabe



J. Am. Chem. Soc. 2020, 142, 8585.