An Unconventional Approach to the Enantioselective Synthesis of Caryophylloids


**Synthesis of (−)-β-Caryophyllene**

**Significance:** The synthesis of β-caryophyllene and coraxenolide A by Larionov and Corey is distinctive because it is a rare example of the use of planar chirality in natural product synthesis. Both enantiomers of (2Z,6E)-6-methylcyclonona-2,6-dienone (H) were prepared and used as chiral precursors for the synthesis of the (−)-β-caryophyllene and coraxenolide A.

**Comment:** The absence of stereoselectivity in the reduction of C with NaNH₄ was overcome by using the CBS reduction. Planar chiral H was obtained as a single enantiomer that is stable against racemization at room temperature owing to restricted C–C bond rotation in the 9-membered ring. By contrast, cyclononene racemizes in a few minutes at room temperature.

**Key words**

- caryophyllene
- planar chirality
- CBS reduction
- Grob fragmentation