## Category

Synthesis of Natural Products and Potential Drugs

## Key words

(±)-paeoniflorigenin

Paeonia lactiflora

Prilezhaev reaction

Reformatsky-type cyclization



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Total Synthesis of (±)-Paeoniflorigenin and Paeoniflorin

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## Synthesis of (±)-Paeoniflorigenin

**Significance:** In 1993, Corey and Wu reported the synthesis of ( $\pm$ )-paeoniflorigenin, a natural product from the roots of *paenia lactiflora*, which are widely used in traditional Chinese medicine. The approach relies on oxidative annulation of dihydro-m-cresol ether **A** and  $Sml_2$ -induced, Reformatsky-type cyclization from an  $\alpha$ -chloro nitrile.

**Comment:** The synthesis commenced with annulation of **A** and **B** to give  $\gamma$ -lactone **C**. Chlorination followed by Prilezhaev epoxidation yields **E**. Cyclization is achieved by treating lactol **F** with an excess of TMS-triflate. Exposure of **H** to Sml<sub>2</sub> leads to formation of the carbon skeleton of paeniflorigenin. Protection of the secondary alcohol followed by reduction of the nitrile gives **K**. Benzoylation and global deprotection completed the total synthesis of ( $\pm$ )-paeniflorigenin in 13 steps from **A**.

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