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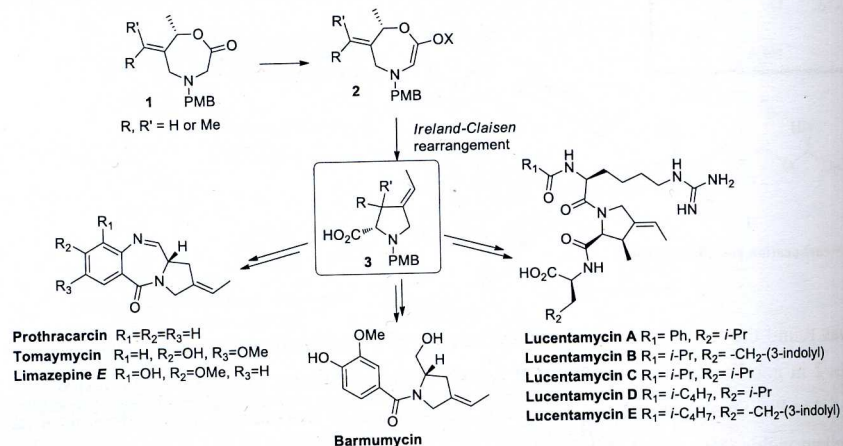
APPLICATION OF IRELAND-CLAISEN REARRANGEMENT IN THE TOTAL SYNTHESIS OF 4-ETHYLIDENE PROLINE CONTAINING NATURAL PRODUCTS

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A number of natural products contain a proline fragment with an *E*-ethylidene substituent at 4 position. Although several total syntheses of these natural products have been reported, efficient control of the olefin geometry has not been achieved.



The key building block for the total syntheses of ethylidene proline containing natural products is proline derivative **3**, possessing all elements of stereochemistry – one or two chiral centers and an *E*-olefin. Herein we disclose an efficient, stereoselective synthesis of ethylidene proline derivatives **3** via a boron enolate Ireland-Claisen rearrangement of 7-membered lactone **1**. The synthesis of 7-membered lactones **1** and the optimization studies of the key transformation – the Ireland-Claisen rearrangement will be presented.

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References:

Smits, G., Zemribo, R. *Org. Lett.* **2013**, *15*, 4406.