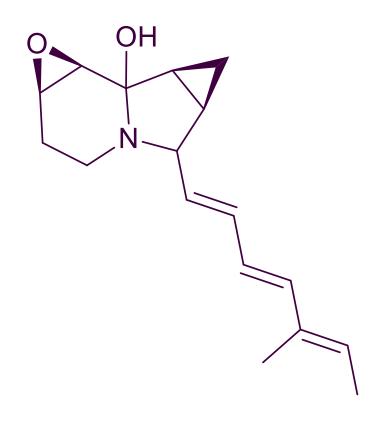
Total Synthesis of Indolizomycin

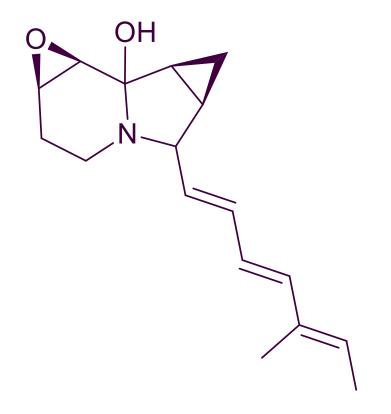
Presented by: Keyvan Pedrood

Instructor: Prof. Robert Maleczka



Spring 2022

Indolizomycin



- A biologically active cyclopropylfused pyrrolidine with remarkable antibiotic activity.
- This compound is produced by a microorganism called SK2-52.
- Total Synthesis of Indolizomycin was proposed by Prof. Samuel J. Danishefsky in 1990.

Retrosynthesis:

Wharton Fragmentation

Julia Coupling

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Protective Group:

Will Be Removed in the Last Step...

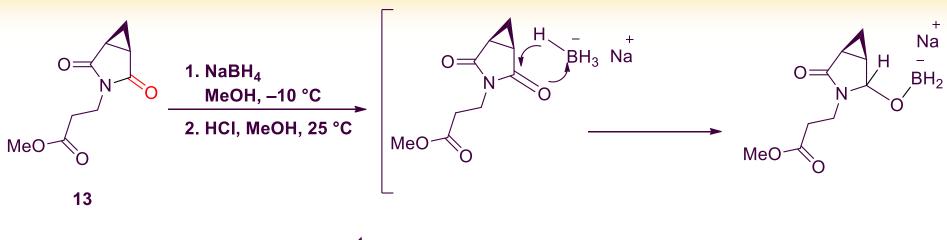
$$\begin{array}{c} Ph_3P \\ N \\ O \\ 16 \end{array}$$

$$\begin{array}{c} O \\ MeO \\ N \\ N \end{array}$$

$$\begin{array}{c} O \\ MeO \\ N \end{array}$$

$$\begin{array}{c} O \\ MeO \\ N \end{array}$$

$$\begin{array}{c} O \\ MeO \\ N \end{array}$$



$$\begin{array}{c} & \\ & \\ -BH_3 \\ -NaCI \end{array}$$

Any Question...?

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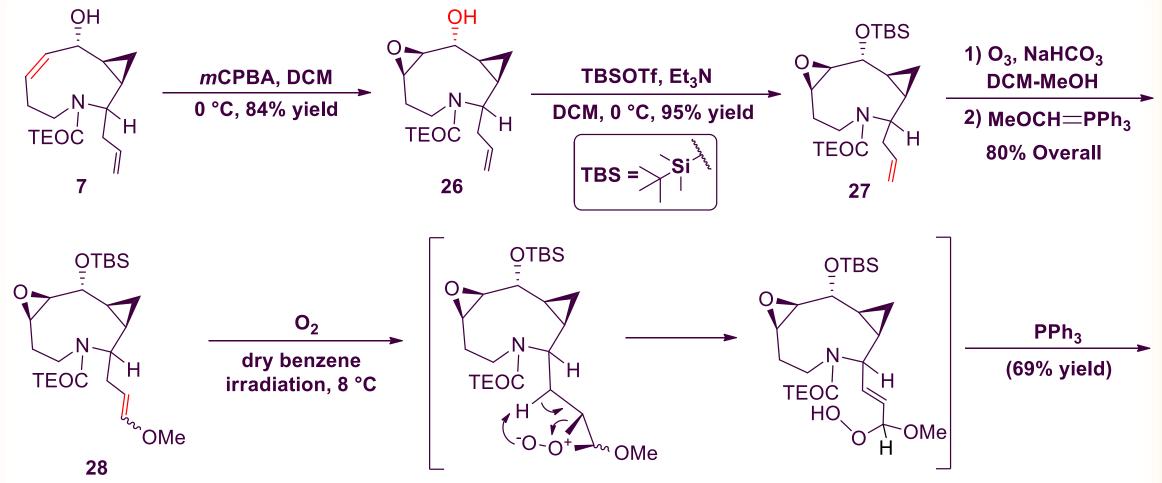
Mechanism:

Any Question...?

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Any Question...?

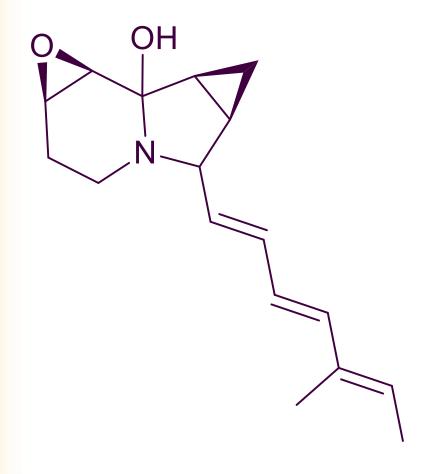
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3:2 mixture of olefin stereoisomers

Any Question...?

Conclusion:



- A number of important name reactions were used to synthesize Indolizomycin including aza-Robinson Annulation, Mcclusky Fragmentation, Wharton Fragmentation, Julia Coupling, Wittig reaction, and Staudinger Reaction.
- Total Synthesis of Indolizomycin by Prof. Samuel J. Danishefsky was, in fact, a masterpiece in contemporary organic synthesis.