



## **Srividhya Sundaram**

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### **Biochemical characterisation of bacterial terpene cyclases**

Terpenes are natural products produced by numerous plants and fungi, traditionally used as flavours and fragrances and are known to possess antibacterial and pharmaceutical properties. Indole terpenoids are derivatives of terpenes containing indole and prenyl moieties and have surprisingly diverse structures catalysed by terpene synthases/cyclases. While most of the information on these compounds and their corresponding enzymes are from eukaryotes, only recently the bacterial candidates have been discovered. Considering the diverse antimicrobial potential of these natural products, a detailed study of their biosynthesis with emphasis on the enzymes becomes necessary.

In my PhD project, I am looking into the biosynthesis of indole terpenoids from *Streptomyces sp.*, and characterize the enzymes critical for the production of these compounds. The major aim of this project is to uncover the reaction mechanism which could be exploited for the synthesis of structurally diverse products. The knowledge of these proteins is fundamental for engineering several indole terpenoids in bacteria and also forms the basis of how these enzymes have evolved to generate natural products with such specificity and fidelity.

## Publications

Sundaram S, Kim HJ, Bauer R, Thongkongkaew T, Heine D, Hertweck C (2018) On-line Polyketide Cyclization into Diverse Medium-Sized Lactones by a Specialized Ketosynthase Domain. *Angew Chem Int Ed Engl* 57(35), 11223-11227. [Details](#) [PubMed](#)

Kugel S, Baunach M, Baer P, Ishida-Ito M, Sundaram S, Xu Z, Groll M, Hertweck C (2017) Cryptic indole hydroxylation by a non-canonical terpenoid cyclase parallels bacterial xenobiotic detoxification. *Nat Commun* 8, 15804. [Details](#) [PubMed](#)

Sundaram S, Hertweck C (2016) On-line enzymatic tailoring of polyketides and peptides in thiotemplate systems. *Curr Opin Chem Biol* 31, 82-94. [Details](#) [PubMed](#)

Heine D, Sundaram S, Bretschneider T, Hertweck C (2015) Twofold polyketide branching by a stereoselective enzymatic Michael addition. *Chem Commun (Camb)* 51(48), 9872-9875. [Details](#) [PubMed](#)

Sundaram S, Heine D, Hertweck C (2015) Polyketide synthase chimeras reveal key role of ketosynthase domain in chain branching. *Nat Chem Biol* 11(12), 949-951. [Details](#) [PubMed](#)

Heine D, Bretschneider T, Sundaram S, Hertweck C (2014) Enzymatic polyketide chain branching to give substituted lactone, lactam, and glutarimide heterocycles. *Angew Chem Int Ed Engl* 53(43), 11645-11649. [Details](#) [PubMed](#)

## Supervisor

[Christian Hertweck](#)

## Start of PhD

July 16, 2012

## Doctoral Disputation

August 11, 2017