

**Ph.D. Open Seminar**  
**Department of Chemistry, IISER Bhopal**

Topic of Seminar: "Synthetic Approaches to Alkaloids Sharing Hexacyclic 3a,3a'-bis-Pyrrolo[2,3-b]indoline Scaffolds: Total Synthesis of (+)- and (-)-Calycanthidine"

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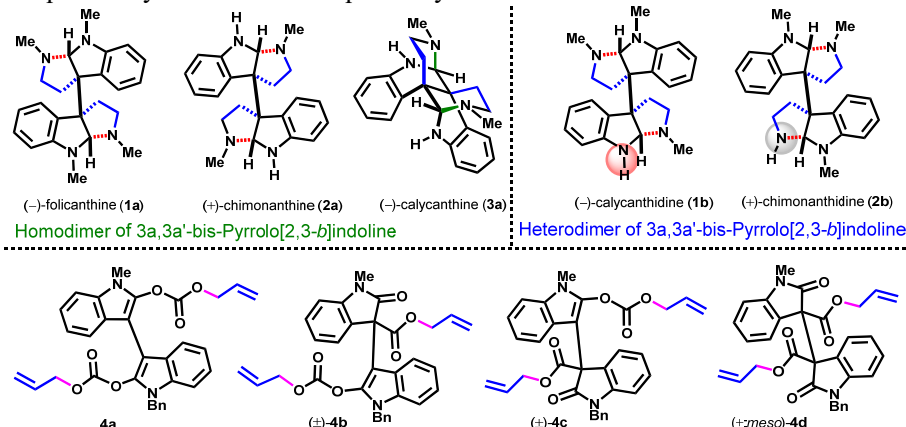
Date: **July 03, 2019**

Time: **4:30 PM**

Venue: **AB2-401**

**Abstract**

A number of naturally occurring dimeric hexahydropyrroloindole (HPI) alkaloids (Figure 1), including homo- and hetero-dimeric scaffolds, exhibit important biological activities such as antifungal and cytostatic properties.<sup>1</sup> Structurally, these molecules usually incorporate a couple of sterically hindered vicinal all-carbon quaternary stereocenters at C3a and C3a', which is regarded as "a daunting challenge" from an organic synthesis perspective.<sup>1a</sup> Due to the steric congestion imposed by the four attached carbons, a very limited number of reports on C-C bond-forming reactions that reliably assemble quaternary carbons are known in literature and hence it puts forward a great synthetic challenge.<sup>1d</sup> The challenge is aggravated even further when two all-carbon quaternary stereocenters are contiguous. Notwithstanding, one of the ways to solve this problem is typically by installing the quaternary stereocenters sequentially.<sup>2-3</sup>



**Figure.** Dimeric indole alkaloids sharing vicinal all-carbon quaternary stereocenters.

During my Ph.D. programme, I undertook synthetic approaches to the alkaloids having hexacyclic 3a,3a'-bis-pyrrolo[2,3-b]indoline ring systems<sup>4a-b</sup> following a highly enantio-, chemo-, and diastereoselective methodology which builds vicinal all-carbon quaternary stereocenters *via* double decarboxylative allylation reactions of compound type 4 to afford products in 92% enantioselectivity with ~17:1 dr (Figure). The strategy has eventually been applied in the total synthesis of (+)- and (-)-calycanthidine.<sup>5</sup>

**References and Notes:**

- (a) Douglas, C. J.; Overman, L. E. *Proc. Natl. Acad. Sci. U.S.A.* **2004**, *101*, 5363. (b) Movassaghi, M.; Schmidt, M. A. *Angew. Chem., Int. Ed.* **2007**, *46*, 3725. (c) Ruiz-Sanchis, P.; Savina, S. A.; Albericio, F.; Álvarez, M. *Chem. - Eur. J.* **2011**, *17*, 1388 and references cited therein.
- (a) Chaudhuri, S.; Bhunia, S.; Roy, A.; Das, M. K.; Bisai, A. *Org. Lett.* **2018**, *20*, 288. (b) Kumar, N.; Maity, A.; Gavit, V. R.; Bisai, A. *Chem. Commun.* **2018**, *54*, 9083. (c) Ghosh, S.; Chaudhuri, S.; Bisai, A. *Chem. - Eur. J.* **2015**, *21*, 17479.
- (a) Babu, N.; Roy, A.; Singh, M.; Bisai, A. *Org. Lett.* **2018**, *20*, 6327. (b) Kumar, N.; Das, M. K.; Ghosh, S.; Bisai, A. *Chem. Commun.* **2017**, *53*, 2170. (c) Babu, N.; Kinthada, L. K.; Das, P. P.; Bisai, A. *Chem. Commun.* **2018**, *54*, 7963. (d) De, S.; Das, M. K.; Roy, A.; Bisai, A. *J. Org. Chem.* **2016**, *81*, 12258.
- Pyrrolo[2,3-b]indoline synthesis under oxidative condition, see; (a) A. Roy, M. K. Das, S. Chaudhuri, A. Bisai, *J. Org. Chem.* **2018**, *83*, 403-421. (c) Roy, A.; Maity, A.; Das, M. K.; Bisai, A. *Manuscript Submitted*.
- (a) Roy, A.; Shaheeda, S.; Babu, K. N.; Bisai, A. *Manuscript Submitted*. (b) Roy, A.; Shaheeda, S.; Maity, A.; *Manuscript under preparation*.