## SYNTHESIS OF MACROCYCLIC COMPOUNDS VIA CONSECUTIVE SONOGASHIRA-UGI-SONOGASHIRA REACTIONS

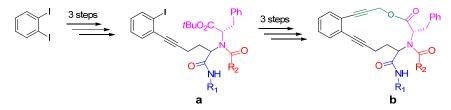
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Multicomponent reactions (MCRs) are among the most important reactions in organic synthesis [1,2]. The main advantages of MCRs are rapid, cost-effective and sustainable access to chemical diverse small-molecule libraries from relatively simple and inexpensive components. The Ugi reaction is the most utilized isocyanide-based multicomponent reactions (IMCRs) which provides a rapid coupling of an aldehyde, an amine, a carboxylic acid and an isocyanide affording  $\alpha$ -acylaminoamides [3].

Compounds with enediyne structural motif were isolated from natural products and showed strong anticancer activity [4]. Apart from their biological profile, enediyne compounds are widely utilized as structural motifs in material chemistry, catalysts design [5], and in metal complexation studies [6].

Our aim was to exploit enediyne structural rigidity in the synthesis of conformationally pre-defined macrocyclic compounds. Our strategy relies on functionalization of 1,2-diiodobenzene [7] and subsequent Sonogashira reaction to yield different Ugi compounds **a**. The second Sonogashira reaction followed by the intramolecular cyclisation afforded highly decorated macrocyclic compounds **b**.



<sup>[1]</sup> G. Koopmanschap, E. Ruijter, R. V. A. Orru, Beilstein J. Org. Chem., 10 (2014) 544 - 598.

<sup>[2]</sup> Brauch, S. S. van Berkel, B. Westermann, Chem. Soc. Rev., 42 (2013) 4948 - 4962.

<sup>[3]</sup> I. Ugi, Angew. Chem. Int. Ed., 1 (1962) 8 - 21.

<sup>[4]</sup> K. C. Nicolaou, W.-M. Dai, Angew. Chem. Int. Ed. 30 (1991) 1387 - 1416.

<sup>[5]</sup> Z. Kokan, Z. Glasovac, M. Majerić Elenkov, M. Gredičak, I. Jerić, S. I. Kirin, Organometallics 2015, 33, 4005.

<sup>[6]</sup> M. Gredičak, N. Bregović, D. Carić, I. Jerić, J. Inorg. Biochem. 2012, 16, 45.

<sup>[7]</sup> M. Glavaš, M. Gredičak, I. Jerić, ACS Comb. Sci. 20 (2018) 151 - 155.