

Supramolecular structuring of cyclo-dipeptides

Joanna Bojarska¹

¹*Technical University*

joanna.bojarska@p.lodz.pl

Cyclo-dipeptides are the simplest cyclic biomolecules that could play a key role in the origin of life [1-2]. They have a huge potential in biomedicine due to their unique features, such as high stability, cell permeability, crossing the blood-brain barrier, stimuli responsiveness, no/low toxicity, structural and biofunctional diversity, easy modification, low costs of production, and so on [3]. They have the inherent ability to build specific highly organized nano-biomimetic structures, through non-covalent interactions, acting as a catalyst to easy gelation. Thus, cyclic dipeptides provide minimalistic scaffolds for self-assemblies, that can have relevance in addressing complex biological problems, inter alia tissue engineering, bio-sensing, or can help to understand biological processes. Notably, the self-assembly process is observed in living organisms, regarding arranging bio-machinery, especially the DNA helix [2].

The design of cyclo-dipeptide-based self-assembled structures still is a challenge. This work provides a deep insight into the supramolecular nature of cyclic dipeptides and considerations on supramolecular structuring of all known, so far, cyclo-dipeptide structures, directed by H-bonding synthon patterns, which can be helpful in the development of innovative drugs, vaccines, diagnostics, drug/gene delivery systems and bio-materials useful in diverse diseases, mainly cancers, viral and neurodegenerative disorders.

1. Bojarska, J.; Mieczkowski, A.; Ziora, Z.M.; Skwarczynski, M.; Toth, I.; Shalash, A.O.; Parang, K.; El-Mowafi, S.A.; Mohammed, E.H.M.; Elnagdy, S.; et al. Cyclic Dipeptides: The Biological and Structural Landscape with Special Focus on the Anti-Cancer Proline-Based Scaffold. *Biomolecules* 2021, 11, 1515.
2. Bojarska, J.; Wolf, W.M. Ultra-short cyclo-peptides as bio-inspired therapeutics: Proline-based 2,5-diketopiperazines (DKP). *Proceedings* 2021, 79, 10.
3. Apostolopoulos, V.; Bojarska, J.; Chai, T.T.; Elnagdy, S.; Kaczmarek, K.; Matsoukas, J.; New, R.; Parang, K.; Lopez, O.P.; Parhiz, H.; et al. A Global Review on Short Peptides: Frontiers and Perspectives. *Molecules* 2021, 26, 430.