DEVELOPMENT SEMIPERMEABLE HOLLOW FIBERSES FOR SEPARATION OF VIOLACEIN PRODUCED IN A MEMBRANE BIOREACTOR

Supervisor: prof. Andrzej Chwojnowski, PhD Eng Co- supervisor: Cezary Wojciechowski PhD Eng achwojnowski@ibib.waw.pl wojciechowski@ibib.waw.pl

Nalecz Institute of Biocybernetics and Biomedical Engineering, PAS Department of Biomaterials and Biotechnological Systems Laboratory of Semipermeable membran es and Bioreactors

The work is to develop a capillary semi-permeable membrane dedicated directly for use in a bioreactor in which bacteria from the strain of *Janthinobacterium lividum* producing violacein will be grown. These membranes are to be used to separate the violacein from the post-culture wort. It will be necessary to develop membranes with specific cut-off points. It is also anticipated that membranes will undergo partial degradation and be intended directly for the bioreactor. The ultimate goal is to isolate from the biosynthesis products the violacein with the highest concentration and highest purity possible to obtain by membrane techniques.

In the era of intensive search for new drugs for one of the most serious multifactorial diseases such as cancer, substances such as violacein are attracting increasing world interest. Violacein is apurple dye produced by some Gram-negative strains. This compound is characterized by a fairly broad spectrum of activity, demonstrating antibacterial, antiparasitic, antifungal, anti-tumor and immunomodulatory properties Interestingly, it seems that the mechanism of violacein action is not universal and is specific for a given cell type. It is also worth noting that violacein can attenuate tumor growth in vivo without inducing adverse effects at higher doses.

It is planned to develop an efficient method of purifying violacein from the strain Janthinobacterium lividum KP16 (strain from the collection of KBŚLiK PW) in the bioproduction process. The strain is currently grown on $\frac{1}{2}$ LB (lysogeny broth) medium at a temperature of about 17 °C with shaking 110-120 rpm. The maximum production of violacein occurs after about 120 hours of breeding.

References:

- Ahmad, W. A., Ahmad, W. Y. W., *i in.* (2012). Isolation of Pigment-Producing Bacteria and Characterization of the Extracted Pigments. Application of Bacterial Pigments as Colorant: The Malaysian Perspective. Berlin, Heidelberg, Springer Berlin Heidelberg: 25-44.
- Alshatwi, A. A., Subash-Babu, P., *i in.* (2016). Violacein induces apoptosis in human breast cancer cells through up regulation of BAX, p53 and down regulation of MDM2. *Exp Toxicol Pathol* **68**(1): 89-97.
- Antonisamy, P. i Ignacimuthu, S. (2010). Immunomodulatory, analgesic and antipyretic effects of violacein isolated from Chromobacterium violaceum. *Phytomedicine* **17**(3-4): 300-4.
- Asencio, G., Lavin, P., *i in.* (2014). Antibacterial activity of the Antarctic bacterium, Janthinobacterium sp. SMN 33.6 against multi-resistant Gram-negative bacteria. *Electronic Journal of Biotechnology* **17**: 1-5.
- Aruldass, C. A., Rubiyatno, *i in*. (2015). Violet pigment production from liquid pineapple waste by Chromobacterium violaceum UTM5 and evaluation of its bioactivity. *RSC Advances* 5(64): 51524-51536.
- Bilsland, E., Tavella, T.A., *i in.* (2018). Antiplasmodial and trypanocidal activity of violacein and deoxyviolacein produced from synthetic operons. *BMC Biotechnol.* **18**(1):22.
- Bromberg, N., Dreyfuss, J. L., *i in*. (2010). Growth inhibition and pro-apoptotic activity of violacein in Ehrlich ascites tumor. *Chem Biol Interact* **186**(1): 43-52.
- Choi, S. Y., Yoon, K. H., *i in*. (2015). Violacein: Properties and Production of a Versatile Bacterial Pigment. *Biomed Res Int* **2015**: 465056.
- Duran, N., Justo, G. Z., *i in*. (2016). Advances in Chromobacterium violaceum and properties of violacein-Its main secondary metabolite: A review. *Biotechnol Adv* **34**(5): 1030-1045.
- Kothari, V., Sharma, S., *i in.* (2017). Recent research advances on Chromobacterium violaceum. *Asian Pac J Trop Med* **10**(8): 744-752.
- Leal, A. M., de Queiroz, J. D., *i in.* (2015). Violacein induces cell death by triggering mitochondrial membrane hyperpolarization in vitro. *BMC Microbiol* **15**: 115-123.
- Verinaud, L., Lopes, S. C., *i in.* (2015). Violacein Treatment Modulates Acute and Chronic Inflammation through the Suppression of Cytokine Production and Induction of Regulatory T Cells. *PLoS One* 10(5): e0125409.
- Woodhams, D.C., LaBumbard, B.C., *i in.* (2018). Prodigiosin, Violacein, and Volatile Organic Compounds Produced by Widespread Cutaneous Bacteria of Amphibians Can Inhibit Two Batrachochytrium Fungal Pathogens. *Microb Ecol* 75:1049-1062.