NON-CONVENTIONAL MEDIA IN FLOW BIOCATALYSIS

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Continuous-flow processing and miniaturization have recently changed the paradigm of biocatalytic process design and significantly contribute to process intensification.^[1] Since many organic substrates or reaction products are only sparingly soluble in water, the use of non-conventional media is receiving considerable attention, especially with the possibility for *in situ* product removal in two-liquid phase systems.^[2] Current trends to use solvents that combine high solubilization capacity and cost-effectiveness with a low environmental footprint, such as deep eutectic solvents and protic ionic liquids that can even improve biocatalyst selectivity, might gain momentum by using high-throughput microflow system for their selection.^[3]

In this presentation, the application of both free enzymes in cosolvent or two-liquid phase systems, as well as immobilized biocatalysts in non-conventional media within microfluidic devices will be highlighted. Examples include lipase-catalyzed esterifications and transesterifications in selected ionic liquids with immobilized enzyme, or in aqueous/organic solvent and ionic liquid/organic solvent systems with dissolved enzyme^[4-7]. Moreover, amine transaminase-catalyzed transamination using selected deep eutectic solvent in a magnetic-field assisted microreactor will be presented, and the effect of various cosolvent and two-liquid phase systems on laccase-catalyzed tyrosol acetate oxidation in a microflow system will be discussed. Finally, the application of deep eutectic solvent for tuning the characteristics of a copolymeric hydrogel used for biocatalyst immobilization will be presented.^[8]

- [1] Žnidaršič-Plazl, P. Biocatalytic process intensification via efficient biocatalyst immobilization, miniaturization, and process integration. *Curr. Opin. Green Sustain. Chem.*, **2021**, 32, 100546.
- [2] Žnidaršič-Plazl, P. Enzymatic microreactors utilizing non-aqueous media. *Chimica Oggi Chem. Today*, 2014, 32: 54-61
- [3] Žnidaršič-Plazl, P. Biotransformations in microflow systems: Bridging the gap between academia and industry. *J Flow Chem.*, **2017**, 7 (3-4): 111-117
- [4] Pohar, A; Plazl, I; Žnidaršič-Plazl, P. Lipase-catalyzed synthesis of isoamyl acetate in an ionic liquid/n-heptane two-phase system at the microreactor scale. *Lab Chip*, **2009**, 9: 3385-3390.
- [5] Pohar, A; Žnidaršič-Plazl, P; Plazl, I. Integrated system of a microbioreactor and a miniaturized continuous separator for enzyme catalyzed reactions. *Chem. Eng. J.* **2012**, 189–190: 376–382.
- [6] Cvjetko, M; Vorkapić-Furač, J; Žnidaršič-Plazl, P. Isoamyl acetate synthesis in imidazolium-based ionic liquids using packed bed enzyme microreactor. *Process Biochem.*, 2012, 47: 1344–1350
- [7] Novak, U; Žnidaršič-Plazl, P. Integrated lipase-catalyzed isoamyl acetate synthesis in a miniaturized system with enzyme and ionic liquid recycle. *Green Proc Synth.*, **2013**, 2: 561-568
- [8] Menegatti, T; Kopač, T; Žnidaršič-Plazl, P. Tuning mechanical characteristics and permeability of alginate hydrogel by polyvinyl alcohol and deep eutectic solvent addition. *Bioengineering*, **2024**, 11 (4), 371