

Università degli Studi di Napoli Federico II

PROJECT TITLE Validation of New Models for Endocrine/Immune Risk Assessment

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Project description (259 words)

Bioactives, additives, and active pharmaceutical ingredients are used by various industries to characterize the marketed product, improve its properties, or facilitate certain stages of production, sometimes without being present in the final product. Some substances, even if allowed by current regulations, are suspected endocrine disruptors and may negatively affect human health by altering the function of the endocrine system. The impact on health is partially known, but uncertainties remain concerning a full risk characterization. The project aims at investigating the toxicity of substances belonging to different chemical classes, used in cosmetics, pharmaceuticals, and/or foods, which are of emerging concern (Contaminants of Emerging Concern, CECs). Some chemicals, contained in personal care products — such as skin lighteners and sunscreens (1) — attracted attention of scientific community due to their potential absorption through the skin, the largest organ of the body. In this project, besides traditional chromatography, biochromatographic techniques and in vitro experiments on cell cultures will be employed. Bio chromatography, using biomimetic stationary phases (sphingomyelin, phosphatidylcholine, cholesterol, and/or proteins), is considered a predictive tool for the crossing of molecules through biological barriers and for their toxicity (2,3,4). At the same time, systems evaluating skin absorption will be used, both traditional skin permeation system (Franz diffusion cells) and through new system such as Permeapad®. The latter features high-performance membranes, and it is already validated for nasal and buccal membrane passage (5,6). Permeapad® has been demonstrated to provide a better reproducibility compared to traditional experiments performed on animal/human skin, therefore playing an important role in the challenge of a more sustainable and ethical research.

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REFERENCES

- 1. Yang F, Yuan T, Ao J, Gao L, Shen Z, Zhou J, Wang B, Pan X. Human exposure risk of organic UV filters: A comprehensive analysis based on primary exposure pathways. *Ecotoxicol Environ Saf.* 2024 Sep 15;283:116800. doi: 10.1016/j.ecoenv.2024.116800.
- 2. Russo G, Piccolo M, Neri I, Ferraro MG, Santamaria R, Grumetto L. Lipophilicity profiling and cell viability assessment of a selected panel of endocrine disruptors. *Chemosphere*. 2023 Feb;313:137569. doi: 10.1016/j.chemosphere.2022.137569.
- 3. Grumetto L, Barbato F, Russo G. Scrutinizing the interactions between bisphenol analogues and plasma proteins: Insights from biomimetic liquid chromatography, molecular docking simulations and in silico predictions. *Environ Toxicol Pharmacol.* 2019 May;68:148-154. doi: 10.1016/j.etap.2019.02.008.
- 4. Russo G, Grumetto L, Barbato F, Vistoli G, Pedretti A. Prediction and mechanism elucidation of analyte retention on phospholipid stationary phases (IAM-HPLC) by in silico calculated physico-chemical descriptors. *Eur J Pharm Sci.* 2017 Mar 1;99:173-184. doi: 10.1016/j.ejps.2016.11.026.
- 5. Bibi HA, Holm R, Bauer Brandl A. Use of PermeaPad for prediction of buccal absorption: A comparison to in vitro, ex vivo and in vivo methods. *European Journal of Pharmaceutical Sciences*. 2016;93:399-404.
- 6. di Cagno M, Bibi HA, Bauer-Brandl A. New biomimetic barrier Permeapad[™] for efficient investigation of passive permeability of drugs. *European Journal of Pharmaceutical Sciences*. 2015;73:29-34.

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