

Short-term Exposure to Perfluorooctanoic Acid Induces Oxidative Stress and Necrotic Cell Death in Human HepG2 Hepatocytes



Katarina Borkovic

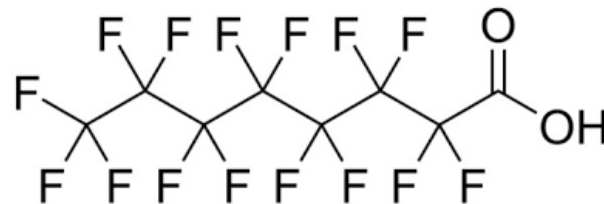
Laboratory for Endocrine
Disruptors and Signaling
(ENDOS)

Dept. of Biology and Ecology
Faculty of Sciences
University of Novi Sad
Novi Sad, Serbia

katarina.borkovic@dbe.uns.ac.rs

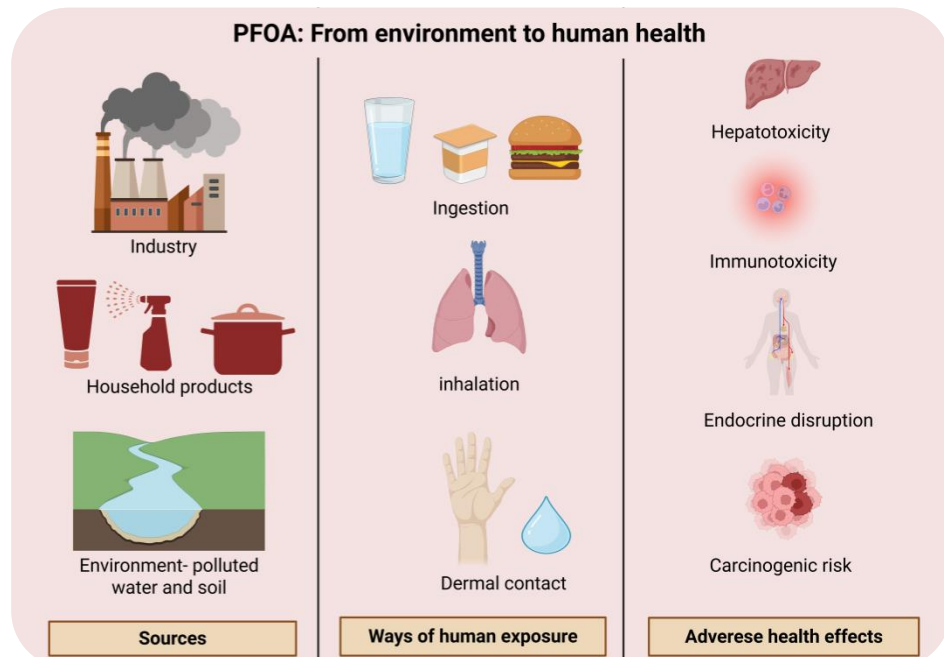
Why look beyond chronic exposure: the case of short-term PFOA exposure

- **Perfluorooctanoic acid (PFOA)** is a synthetic chemical belonging to the class of PFAS, a highly environmentally persistent and bioaccumulative group of substances

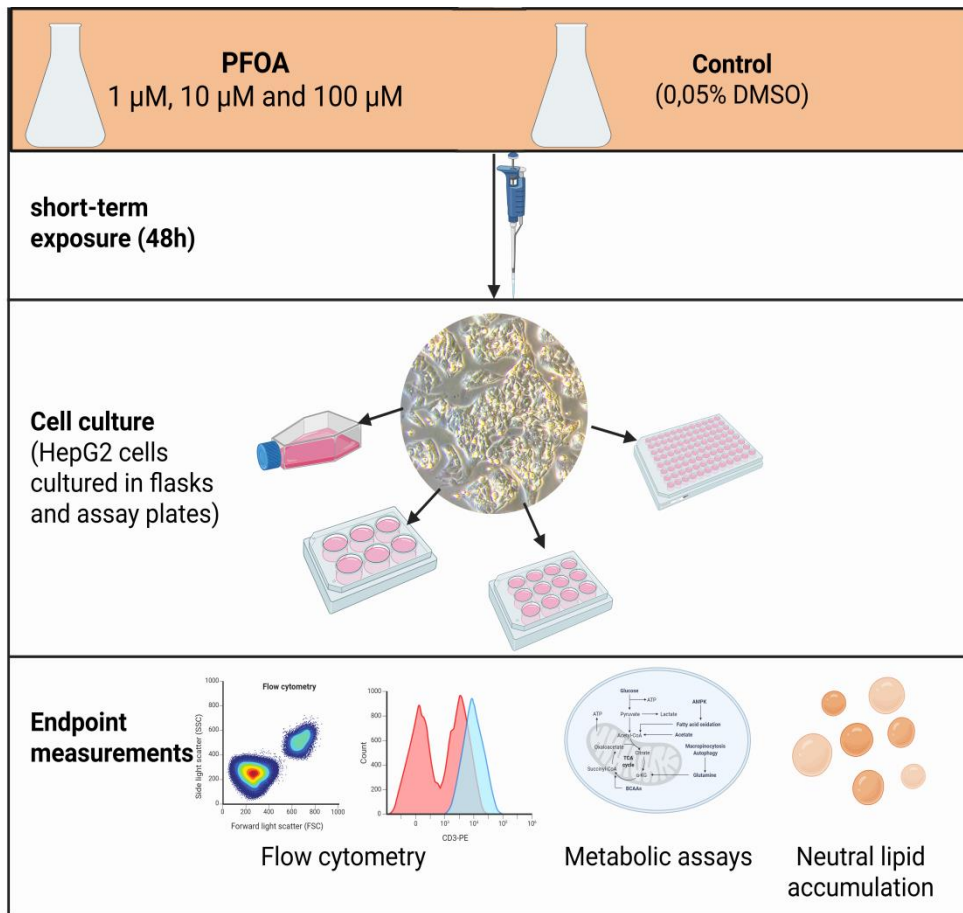


Structure of perfluorooctanoic acid (PFOA)

- Resistant to degradation, accumulates in human tissues
- Linked to immunotoxicity, hepatotoxicity, endocrine disruption, and carcinogenic risk
- Acute cellular effects remain unclear



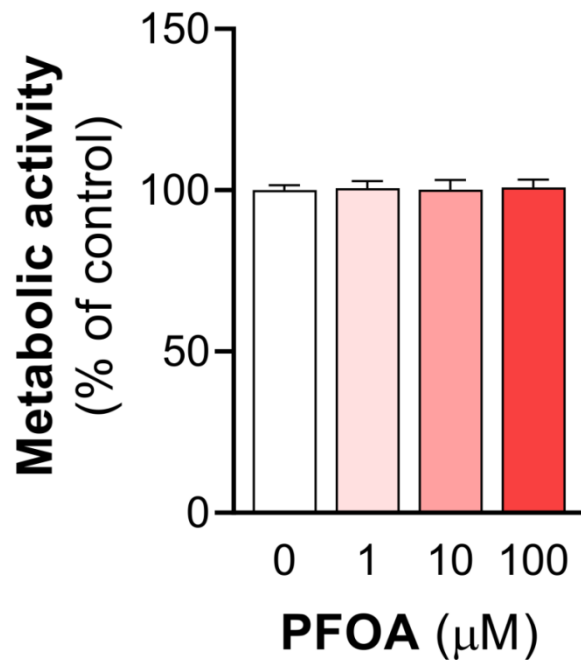
The Pipeline



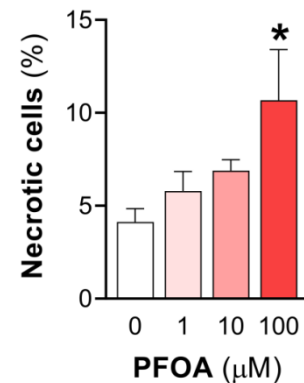
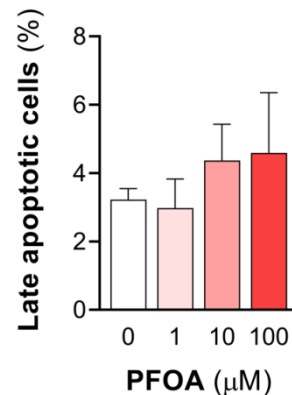
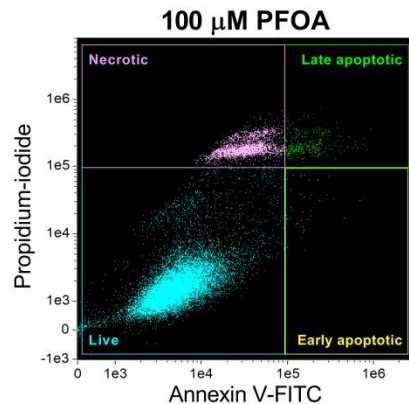
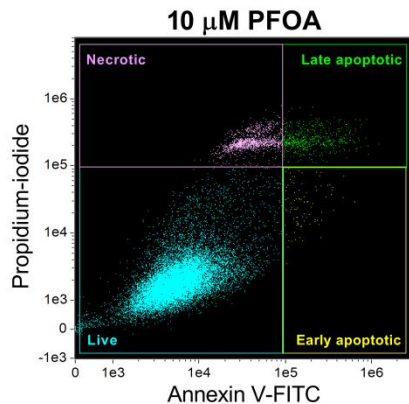
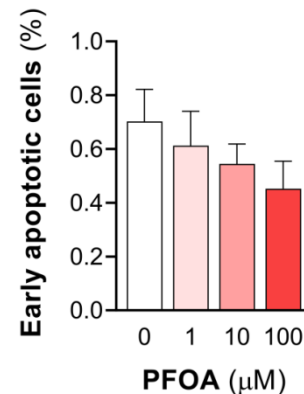
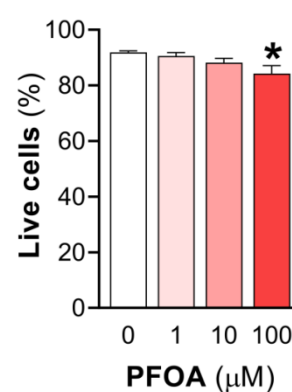
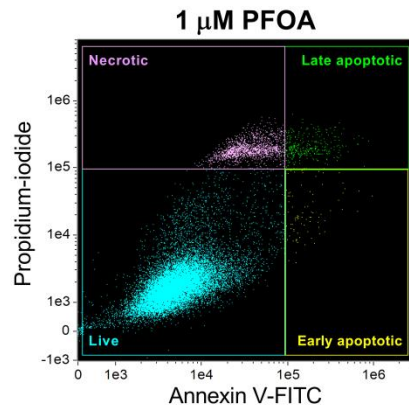
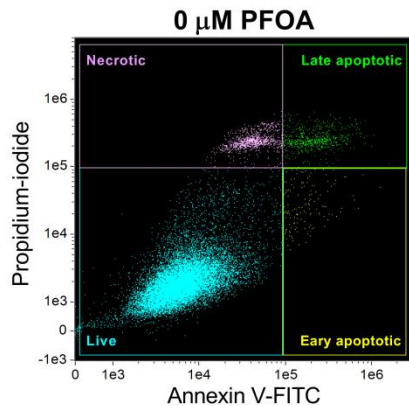
Assessed endpoints:

- **Metabolic activity** – alamarBlue™ assay
- **Cell death pathways** – Annexin V/propidium iodide flow cytometry
- **Cell cycle distribution** – Propidium iodide flow cytometry
- **Intracellular reactive oxygen species (ROS)** – Dichlorofluorescein fluorescence
- **Lipid peroxidation** – Thiobarbituric acid reactive substances (TBARS) assay
- **Neutral lipid accumulation** – Oil Red O staining

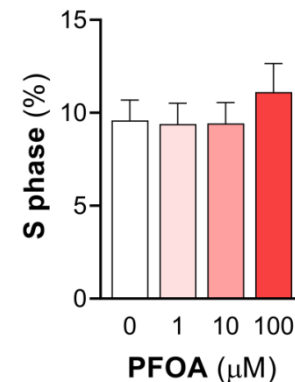
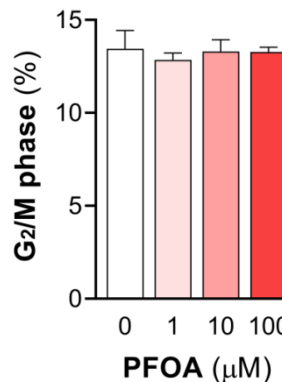
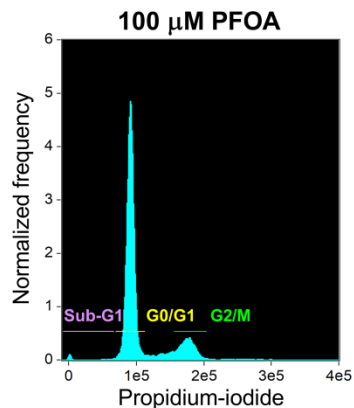
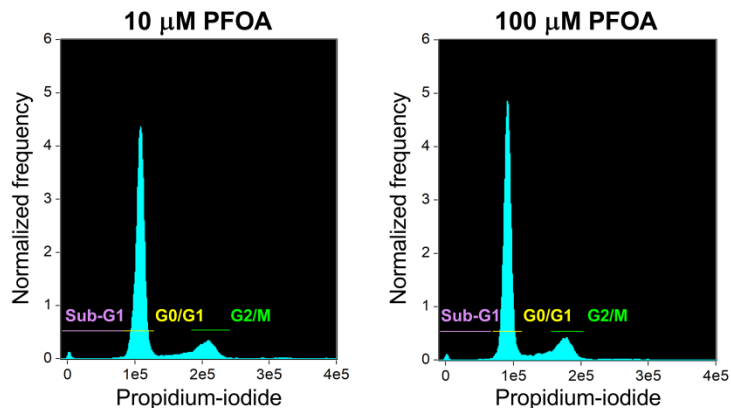
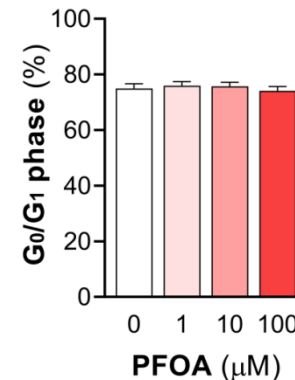
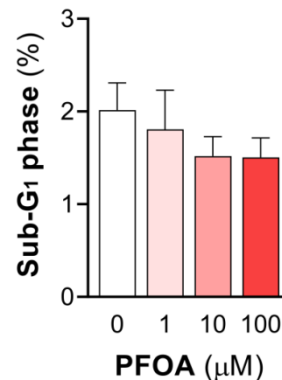
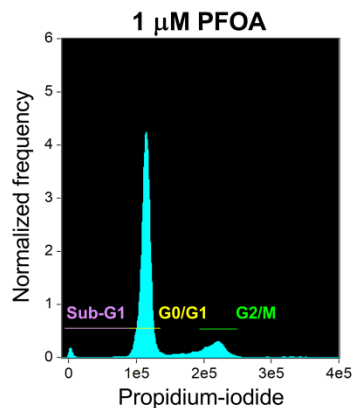
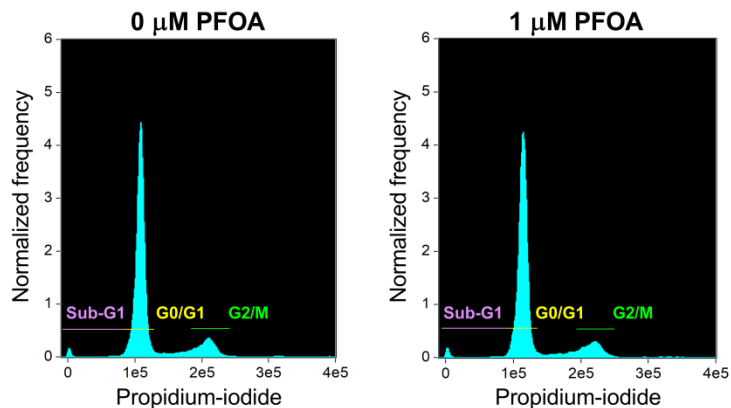
Short-term PFOA exposure does not affect metabolic activity in HepG2 cells



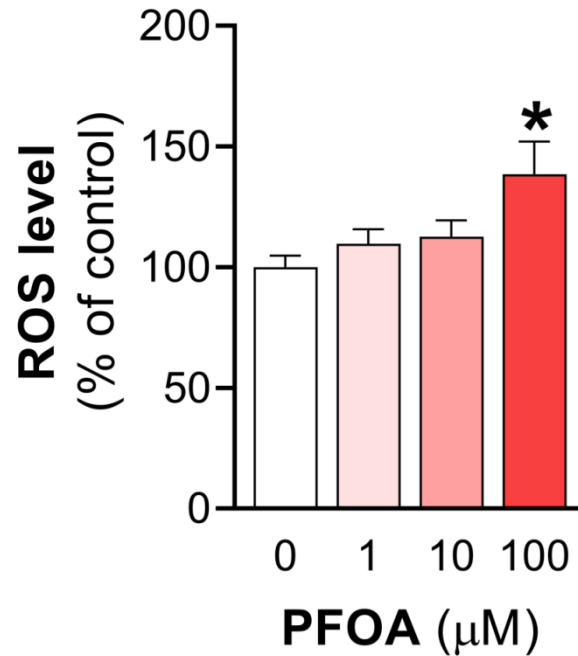
Short-term PFOA exposure reduces cell viability and promotes necrosis in HepG2 cells



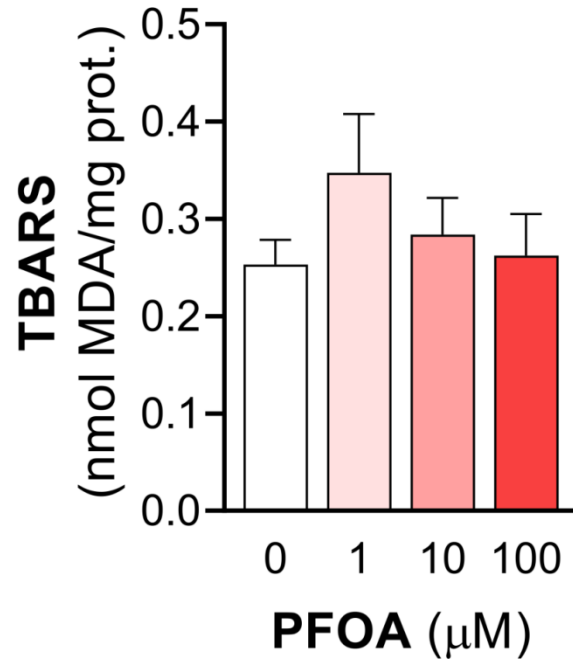
Short-term PFOA exposure does not affect cell cycle progression in HepG2 cells



Short-term PFOA exposure elevates intracellular ROS levels in HepG2 cells

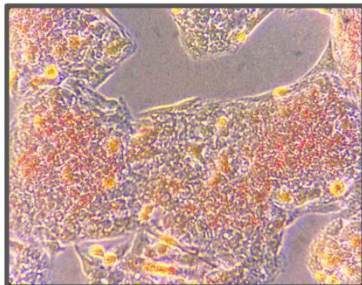


Short-term PFOA exposure does not increase lipid peroxidation in HepG2 cells

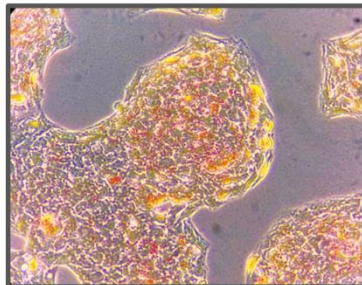


Short-term PFOA exposure does not affect neutral lipid accumulation in HepG2 cells

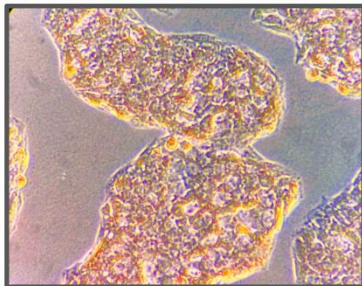
0 μM PFOA



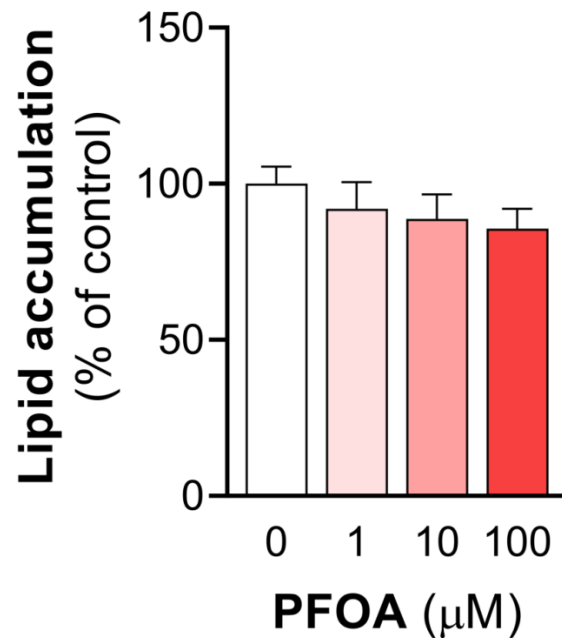
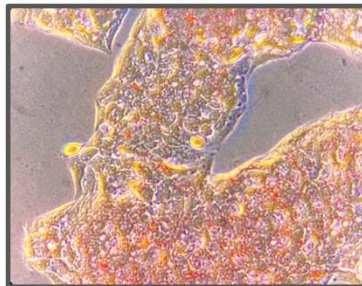
1 μM PFOA



10 μM PFOA

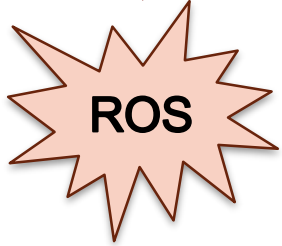


100 μM PFOA



Key Findings

Short-term PFOA exposure induces oxidative stress



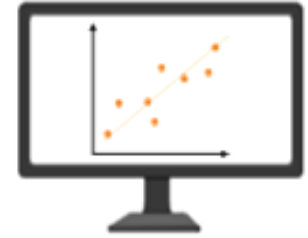
Cell fate shifts toward necrosis rather than apoptosis



No immediate disruption of lipid metabolism or proliferation observed



Provides a snapshot of acute toxicity and baseline for future long-term studies



Acknowledgements:

ENDOS Laboratory

Dunja Kokai

Bojana Stanic

Nebojsa Andric

This study was supported by Science Fund of the Republic of Serbia, #7010, Integration of Biological Responses and PBTK Modeling in Chemical Toxicity Assessment: A Case Study of Perfluorooctanoic Acid (PFOA) – **ToxIN**



Science Fund
of the Republic of Serbia



Katarina Borkovic

Laboratory for Endocrine Disruptors and Signaling (ENDOS)

Dept. of Biology and Ecology
Faculty of Sciences
University of Novi Sad
Novi Sad, Serbia

katarina.borkovic@dbe.uns.ac.rs