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Title of diploma thesis: Cytotoxicity testing on 2D and 3D model of human liver cells.

An inherent part of drug development are *in vitro* assays, which might be helpful in prediction of drug toxicity. Nowadays, the majority of assays use simple 2D structures for cell growth, but 3D structures with similar conditions to in vivo are becoming more popular. The goal of the study was to assess the cytotoxicity of selected xenobiotics in vitro by both 2D and 3D cell models. The research subjects were drugs from the group of antimycotics (amphotericin B, ketoconazole), NSAIDs (diclofenac, ibuprofen), antipyretics (paracetamol, fenacetine), sodium azide, tamoxifen, para-aminosalicylic acid, methanol and ethanol. For determination of cytotoxicity, the standard colorimetric method (CellTiter 96<sup>®</sup>) based on reductive assessment of metabolic active cells was used. For drug testing it was used human standard line of liver cells HepG2. The cells were cultivated in monolayer or in 3D form with the Alvetex<sup>®</sup>Scaffold technology using high porous networked polystyrene. The parameter of inhibition concentration IC50 was chosen for toxicity assessment of tested drugs and it enabled the comparison of drugs cytotoxicity based on 2D and 3D cultures and between each other. Measured values of IC<sub>50</sub> show that cells cultivated in 3D conditions, which are more similar to in vivo conditions, were more resistant against higher concentrations of tested drugs compared to 2D conditions. Amphotericin B showed the highest cytotoxicity in both 2D and 3D model HepG2 cells and phenacetine showed the lowest cytotoxicity in the same conditions. The cytotoxicity effect of phenacetine was relatively low, and therefore, the certain value of IC<sub>50</sub> was not possible to determine, and it was likely occurring out the interface of tested concentrations. With CellTiter  $96^{\text{®}}$  assay it was possible to gain IC<sub>50</sub> for almost all tested drugs and it enabled us to compare cytotoxicity of substances in 2D and 3D models of HepG2 cells. As a conclusion of the findings of in vitro assays the 3D model represents more suitable tool for prediction of drugs induced toxicity.