



Erratum to Body-in-a-Cube: a microphysiological system for multi-tissue co-culture with near-physiological amounts of blood surrogate

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This article that appeared in: Vol 4 (June 2020) Issue of the *Microphysiological Systems (MPS)* (1), was miscategorized as review article. The content of the article justify it as an Original Article. For correction, the abstract of the paper should be replaced by a structured abstract and the correct abstract is presented below:

Background: Decreasing the amount of liquid inside microphysiological systems (MPS) can help uncover the presence of toxic drug metabolites. However, maintaining near-physiological volume ratios among blood surrogate and multiple organ mimics is technically challenging. Here, we developed a body cube and tested its ability to support four human tissues (kidney, GI tract, liver, and bone marrow) scaled down from in vivo functional volumes by a factor of 73,000 with 80 μ L of cell culture medium (corresponding to $\sim 1/73,000$ th of in vivo blood volume).

Methods: GI tract cells (Caco-2), liver cells (HepG2/C3A), bone marrow cells (Meg-01), and kidney cells (HK-2) were co-cultured inside the body cube with 80 μ L of common, recirculating cell culture medium for 72 h. The system was challenged with acetaminophen and troglitazone, and concentrations of aspartate aminotransferase (AST), albumin, and urea were monitored over time.

Results: Cell viability analysis showed that $95.5\% \pm 3.2\%$ of liver cells, $89.8\% \pm 4.7\%$ of bone marrow cells, $82.8\% \pm 8.1\%$ of GI tract cells, and $80.1\% \pm 11.5\%$ of kidney cells were viable in co-culture for 72 h. Both acetaminophen and troglitazone significantly lowered cell viability in the liver chamber as indicated by viability analysis and a temporary increase of AST in the cell culture medium. Both drugs also lowered urea production in the liver by up to 45%.

Conclusions: Cell viability data and the production of urea and albumin indicate that the co-culture of GI tract, liver, bone marrow, and kidney tissues with near-physiological volume ratios of tissues to blood surrogate is possible for up to 72 h. The body-cube was capable of reproducing liver toxicity to HepG2/C3A liver cells via acetaminophen and troglitazone. The developed design provides a viable format for acute toxicity testing with near-physiological blood surrogate to tissue volume ratios.

Keywords: Multi-organ microphysiological system; microphysiological systems (MPS); body-on-a-chip; body cube; microfluidic cell culture

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References

1. Chen L, Yang Y, Ueno H, et al. Body-in-a-Cube: a microphysiological system for multi-tissue co-culture with near-physiological amounts of blood surrogate. *Microphysiol Syst* 2020;4:1.

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