

## Supporting Information

### Development of a biomimetic liver tumor-on-a-chip model based on decellularized liver matrix for toxicity testing

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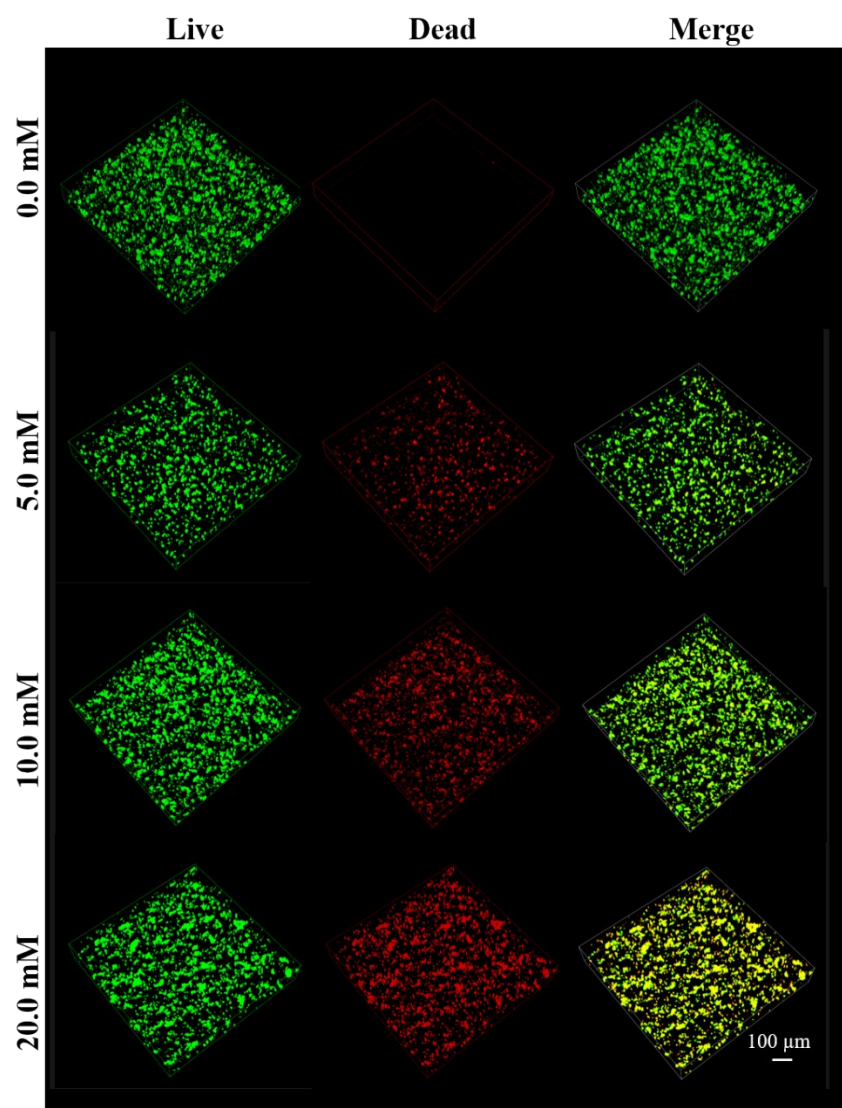
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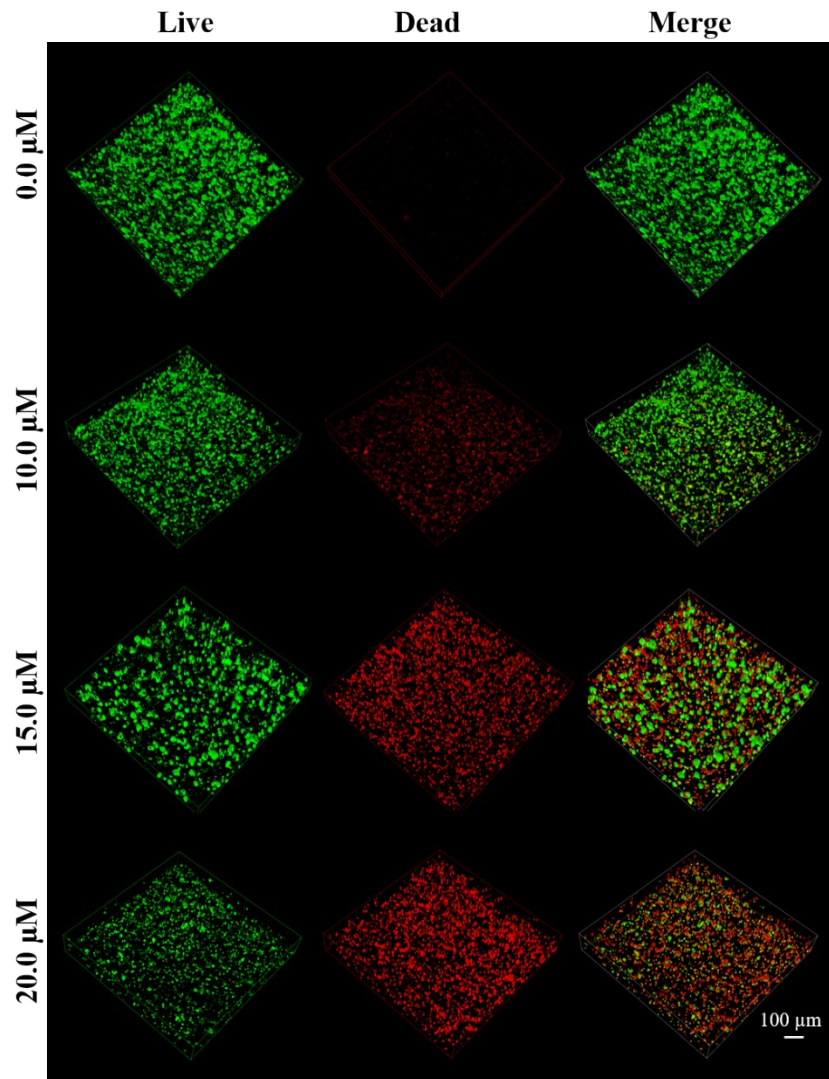
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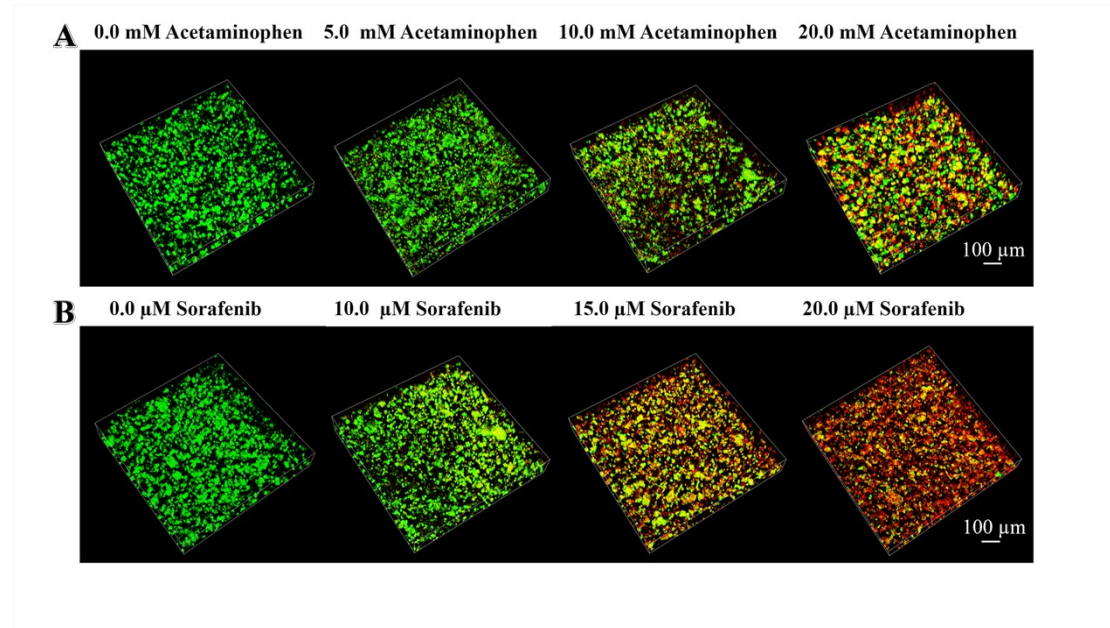
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**Fig. S1** Live/dead staining of HepG2 cells in the 2:3 DLM-GelMA hydrogels after adding 0.0, 5.0, 10.0, and 20.0 mM of acetaminophen after 24h. Live cells were stained in green, and dead cells are stained in red. The yellow color represented the overlaid images of live and dead cells.



**Fig. S2** Live/dead staining of HepG2 cells in the 2:3 DLM-GelMA hydrogels after adding 0.0, 10.0, 15.0, and 20.0  $\mu\text{M}$  of sorafenib after 24h. Live cells were stained in green, and dead cells are stained in red. The yellow color represented the overlaid images of live and dead cells.



**Fig. S3** Evaluation of acetaminophen/sorafenib induced toxicity using the tumor-on-a-chip platform with embedded 3D HepG2-laden GelMA. (A) Live/dead staining showing acetaminophen (0.0, 5.0, 10.0 and 20.0 mM) treatment of HepG2 embedded in GelMA for 24 h. Scale bar: 100  $\mu\text{m}$ . (B) Live/dead staining showing sorafenib (0.0, 10.0, 15.0 and 20.0  $\mu\text{M}$ ) treatment of HepG2 embedded in GelMA only for 24 h. Scale bar: 100  $\mu\text{m}$ .

**Table S1.** Calculation of collagen concentration in native liver, DLM solution and DLM-UV light.

	Native liver	DLM solution	DLM-UV light
Sample Volume ( $\mu\text{L}$ )	400	400	400
HYP ( $\mu\text{g}$ )	3.480 $\pm$ 0.586	8.018 $\pm$ 2.819	8.213 $\pm$ 1.317
Collagen ( $\mu\text{g}$ )	25.058 $\pm$ 4.22	57.731 $\pm$ 20.297	59.135 $\pm$ 9.486
Dry weight (mg)	3	0.4	0.4
Dry Weight Collagen ( $\mu\text{g}/\text{mg}$ )	8.35 $\pm$ 2.44	144.33 $\pm$ 87.89	147.84 $\pm$ 41.07
Total Collagen in the sample ( $\mu\text{g}$ )	25.1	57.7	59.1
Concentration of Collagen in live tissues ( $\mu\text{g}/\mu\text{L}$ )	0.06275	0.14433	0.14784
Concentration of Collagen from DLM in Chip ( $\mu\text{g}/\mu\text{L}$ )	N/A	N/A	0.0591

**Table S2.** Preparation of different volume ratios of DLM-GelMA

DLM-GelMA	DLM ( $\mu\text{L}$ )	10 wt% GelMA ( $\mu\text{L}$ )	PI (g)
GelMA only	0	1,000	0.01
1:9 DLM-GelMA	100	900	0.01
1:4 DLM-GelMA	200	800	0.01
3:7 DLM-GelMA	300	700	0.01
2:3 DLM-GelMA	400	600	0.01
1: 1 DLM-GelMA	500	500	0.01

Abbreviations: PI = photoinitiator

**Table S3.** Dimensions of liver tumor-on-a-chip

Dimensions	Top Layer	Bottom Layer
Width of chamber ( $w$ )	5 mm	5 mm
Length of chamber ( $l$ )	15 mm	15 mm
Height of chamber ( $h$ )	0.5 mm	1 mm
Volume	45 mm <sup>3</sup>	60 mm <sup>3</sup>
Diameter of microchannel	1.5 mm	1.5 mm
Inner diameter of tubes	0.5 mm	0.5 mm
Outer diameter of tubes and of device inlets/outlets	2 mm	2 mm

**Table S4.** Quantification of collagen, glycosaminoglycans (GAG), HGF and bFGF in the DLM solution before and after UV light exposure and in the native liver.

Parameter	Native liver	DLM solution	DLM-UV light
Collagen ( $\mu\text{g}/\text{mg}$ dry weight)	$8.35 \pm 1.41$	$144.33 \pm 50.74$	$147.84 \pm 23.71$
GAG ( $\mu\text{g}/\text{mg}$ dry weight)	$195.23 \pm 5.33$	$499.16 \pm 14.02$	$479.42 \pm 28.77$
HGF ( $\text{pg}/\text{mg}$ dry weight)	$6.70 \pm 0.08$	$4.27 \pm 0.05$	$4.37 \pm 0.04$
bFGF ( $\text{pg}/\text{mg}$ dry weight)	$5.29 \pm 0.09$	$4.58 \pm 0.11$	$4.07 \pm 0.02$

Abbreviations: GAG = glycosaminoglycan; HGF = hepatocyte growth factor; bFGF = basic fibroblast growth factor

**Table S5.** Measurement of Young's modulus and pore size for different volume ratios of DLM-GelMA

	Young's modulus (kPa)	Pore size ( $\mu\text{m}$ )
GelMA only	$19.56 \pm 1.30$	$41.28 \pm 1.79$
1:9 DLM-GelMA	$21.78 \pm 0.45$	$56.74 \pm 5.91$
1:4 DLM-GelMA	$19.41 \pm 0.73$	$72.07 \pm 4.81$
3:7 DLM-GelMA	$19.47 \pm 1.45$	$89.87 \pm 6.27$
2:3 DLM-GelMA	$16.91 \pm 0.54$	$100.34 \pm 8.56$
1:1 DLM-GelMA	$19.14 \pm 0.70$	$101.54 \pm 5.97$