

## A vascularized organ-on-a-chip platform for large-scale drug screening applications

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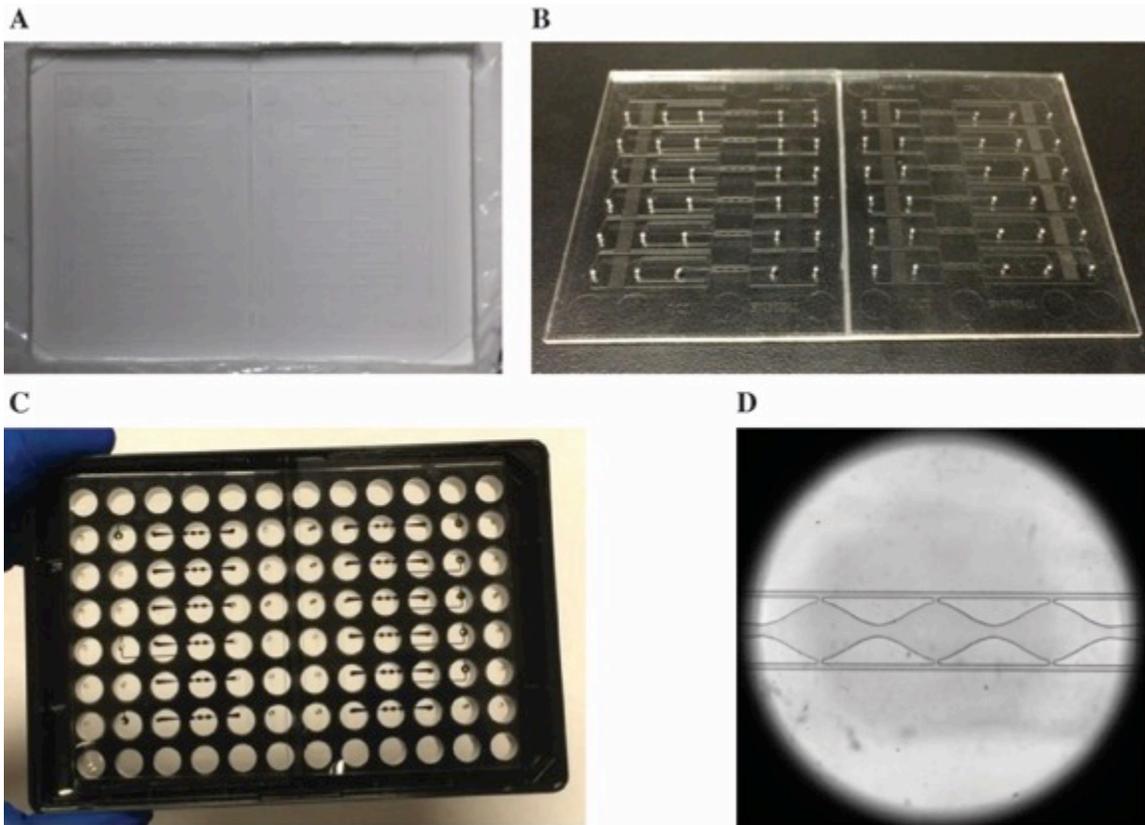
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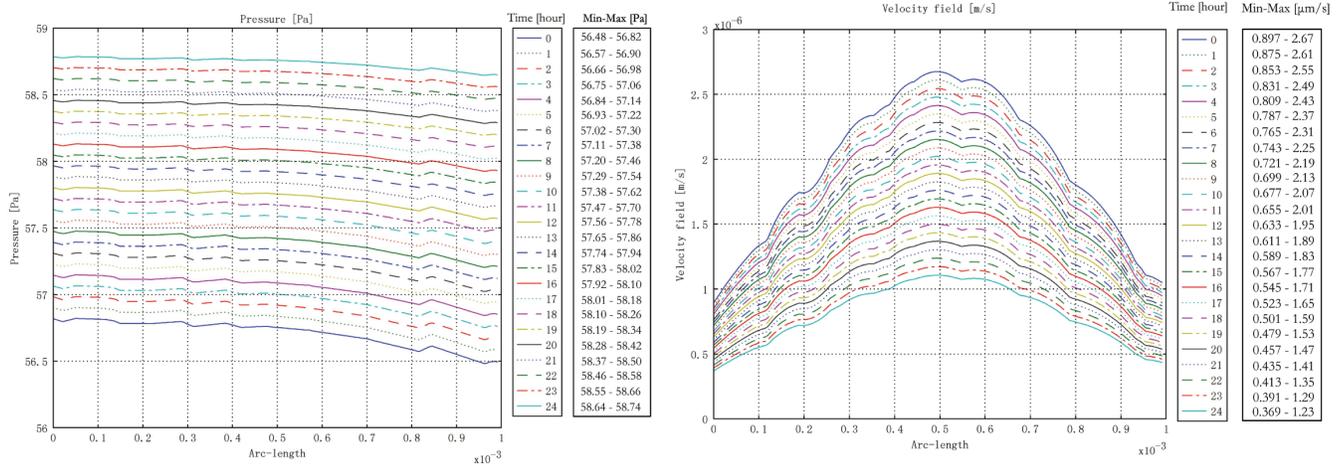
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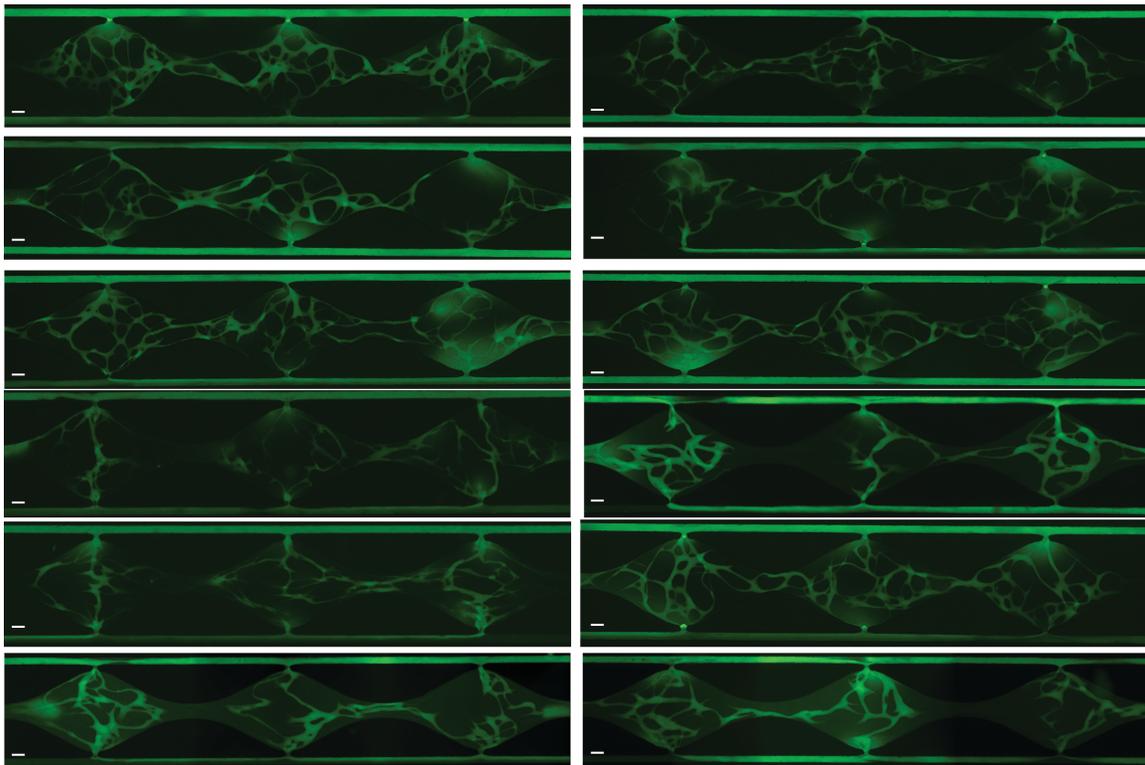
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**Supplemental Figure S1:** Platform fabrication. (A) A customized master mold is fabricated using 2-part polyurethane liquid plastic and a micro-molding technique. (B) A PDMS device layer is replicated from the polyurethane master mold, and holes are punched for inlets and outlets. (C) A bottom view of a fully assembled platform. (D) A low-power view of 3 tissue chambers inside a single well.

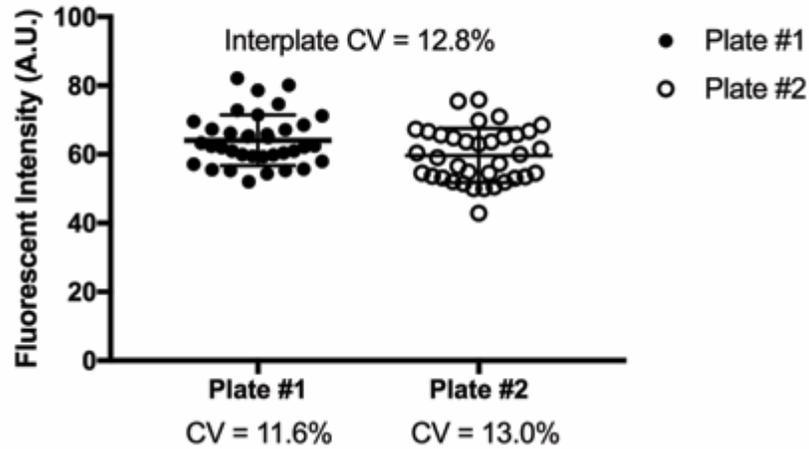


**Supplemental Figure S2:** Finite element simulation of hydrostatic pressure and interstitial velocity flow in horizontal direction of a tissue chamber.

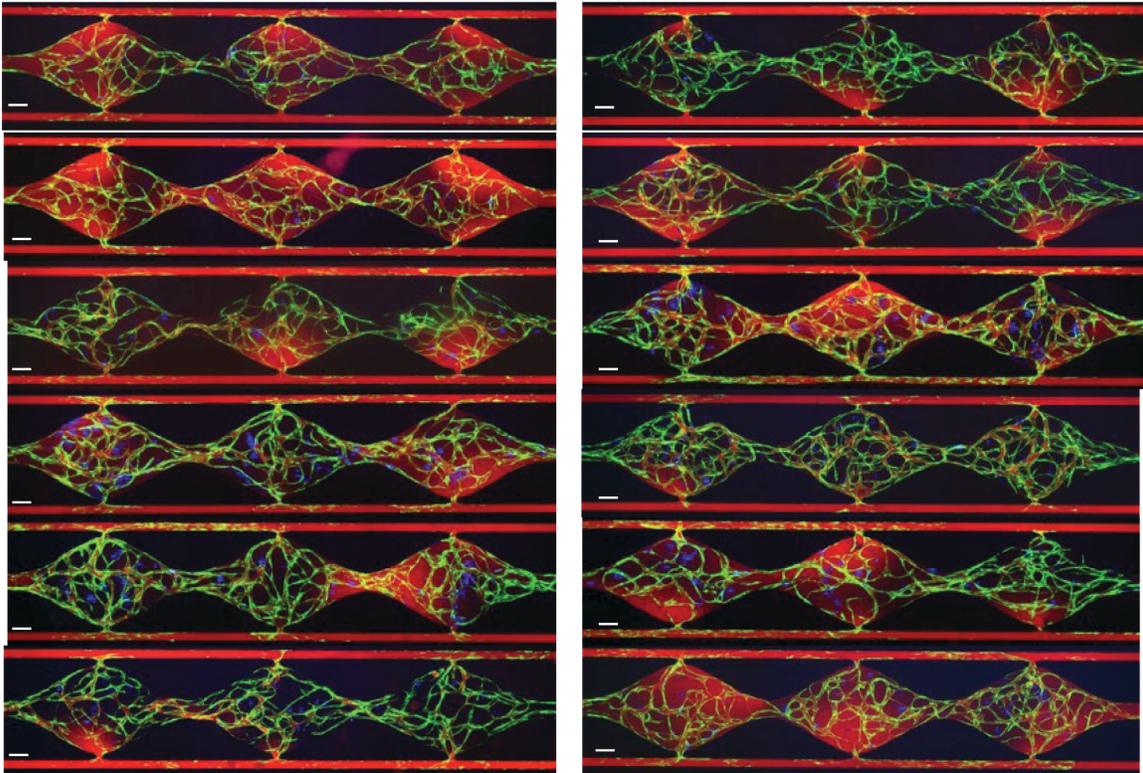


**Supplemental Figure S3:** 70 kDa-FITC dextran perfusion in 12 tissue units within a single platform. Scale bar = 100 µm.

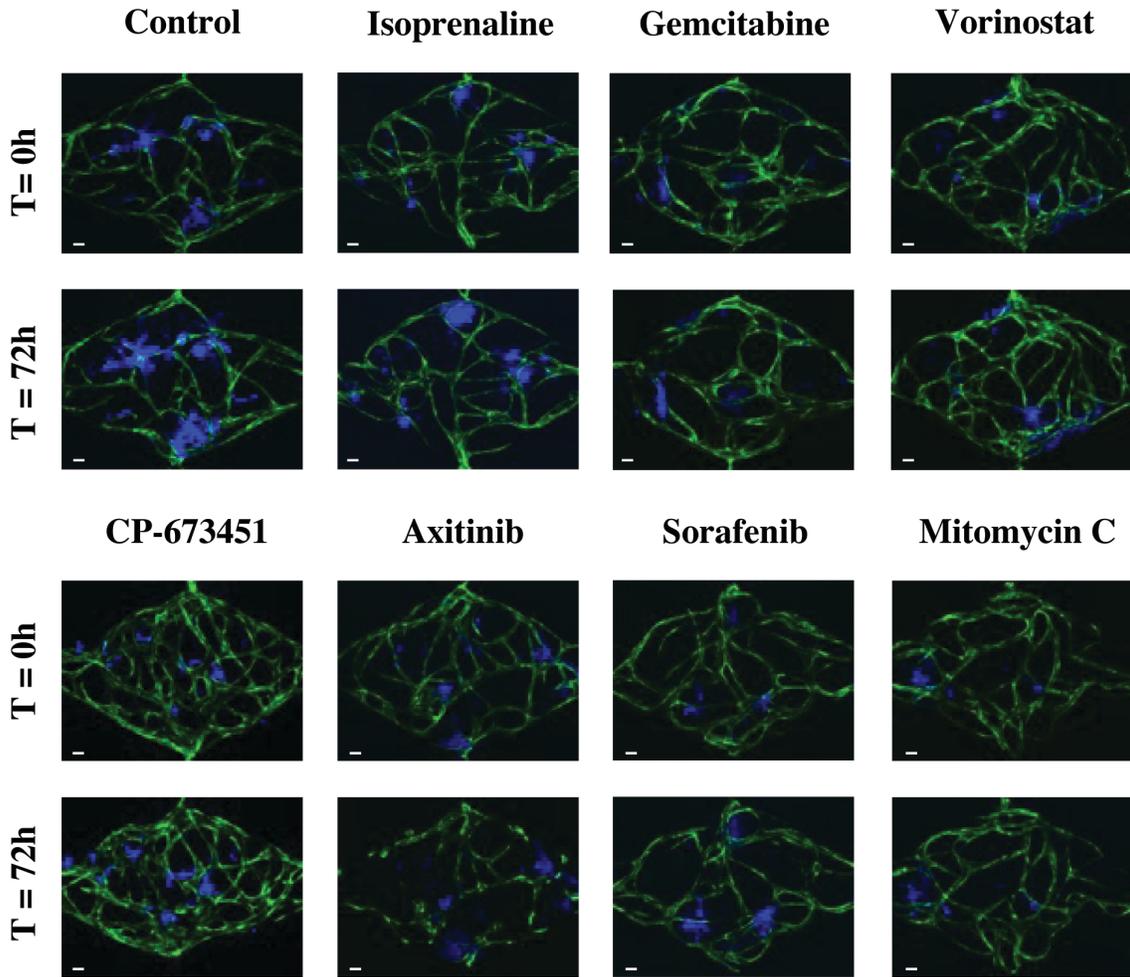
### Fluorescent intensity of monolayer EC in 96-well plate



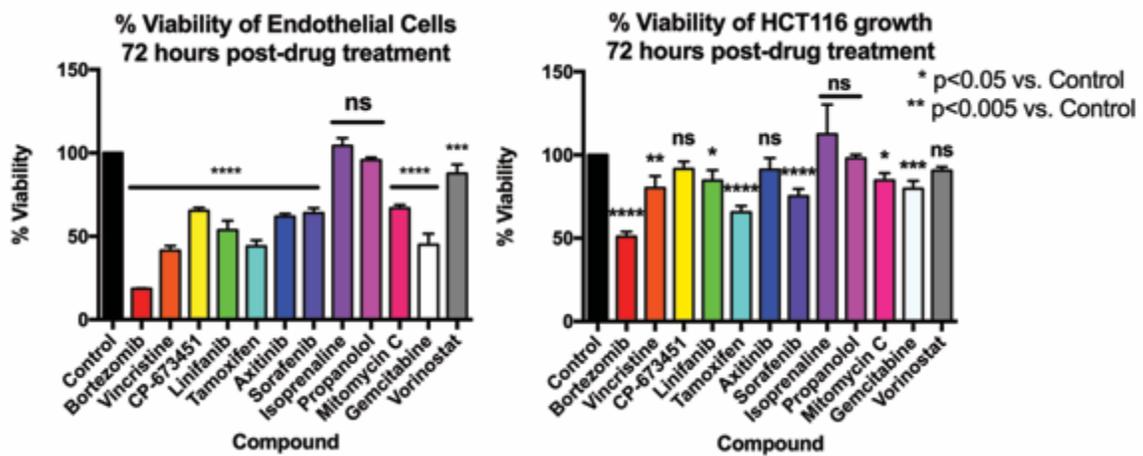
**Supplemental Figure S4:** Coefficient of variation (CV) of a standard 2D monoculture assay. Fluorescent-tagged endothelial cells were plated in 2 independent 96-well plates and allowed to adhere for 2 hours. Fluorescent intensity in each well was measured using a fluorescent plate reader and used to calculate the CV.



**Supplemental Figure S5:** 70 kDa-Rhodamine B dextran perfusion in 12 VMTs within a single platform. Scale bar = 100  $\mu$ m.



**Supplemental Figure S6:** Representative images before (T= 0h) and after (T= 72h) of drug treatment in the VMTs. Scale bar = 50  $\mu$ m.



**Supplemental Figure S7:** Blinded, primary drug screening at 1  $\mu$ M in 2D monoculture assay. Cell viability was quantified using XTT assay after 72 hours of drug treatment.

**Supplemental Table S1:** Sequence of qRT-PCR primers.

Gene Name	Sequence 5'→3'	
<b>18S</b>	CCCCGGCCGTCCTCTTA	Forward
	CGCCCCCTCGATGCTCTTAG	Reverse
<b>ICAM-1</b>	CAGAGGTTGAACCCCCACAGT	Forward
	TCTGAGACCTCTGGCTTCGT	Reverse
<b>E-selectin</b>	CCGTCCGCCAGCCTCAGAAT	Forward
	TAGCCTCGCTCGGGGTTGGAC	Reverse
<b>VCAM-1</b>	CCATTTGACAGGCTGGAGAT	Forward
	TACTGTGGGCAGAGAATCCA	Reverse

**Supplemental Table S2:** Summary of drug compounds and their targets.

Compound	Target(s)
Bortezomib	20S Proteasome
Vincristine	Microtubule inhibitor
CP-673451	Multi-tyrosine kinase inhibitor (Concentration $\leq 1 \mu\text{M}$ : PDGFR- $\alpha/\beta$ , c-Kit, VEGFR1/2)
Linifanib	Potent ATP-competitive VEGFR/PDGFR inhibitor
Tamoxifen	Estrogen receptor antagonist
Axitinib	Multi-tyrosine kinase inhibitor (Concentration $\leq 1 \mu\text{M}$ : VEGFR1/2/3, PDGFR- $\alpha/\beta$ )
Sorafenib	Multi-tyrosine kinase inhibitor (Concentration $\leq 1 \mu\text{M}$ : VEGFR2, Raf, PDGFR- $\beta$ , c-Kit, FGFR1)
Isoprenaline	$\beta$ -adrenergic receptor agonist
Propranolol	$\beta$ -adrenergic receptor inhibitor
Mitomycin C	DNA synthesis inhibitor
Gemcitabine	DNA synthesis inhibitor
Vorinostat	HDAC inhibitor

**Source:** Selleck Chemicals.