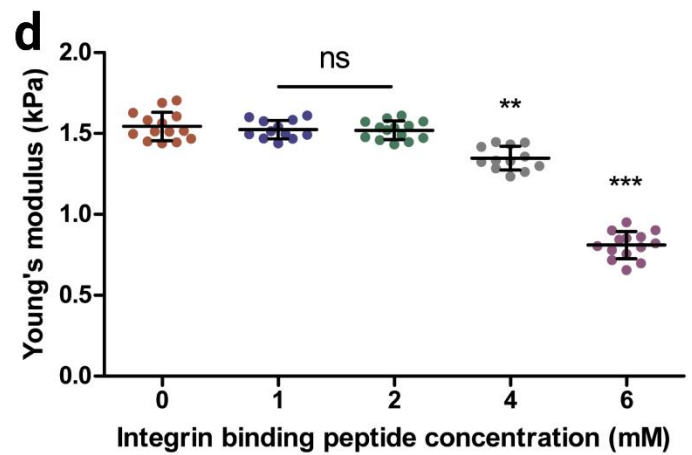
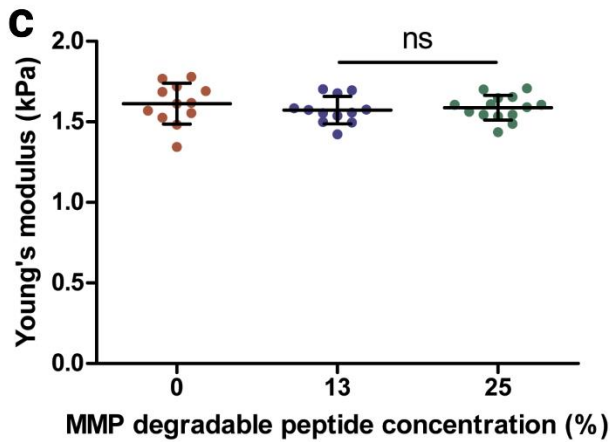
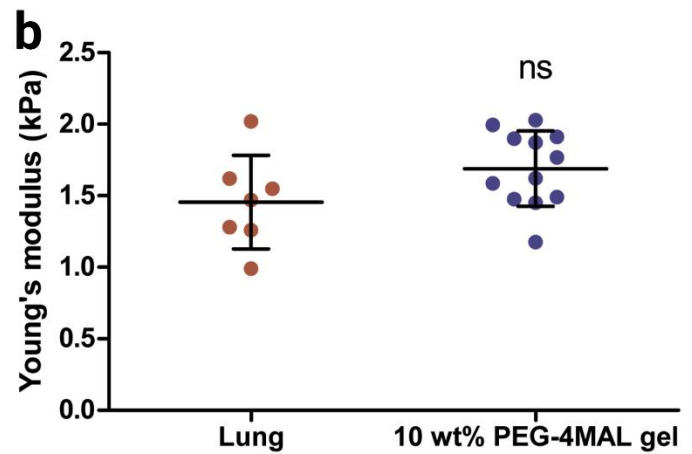
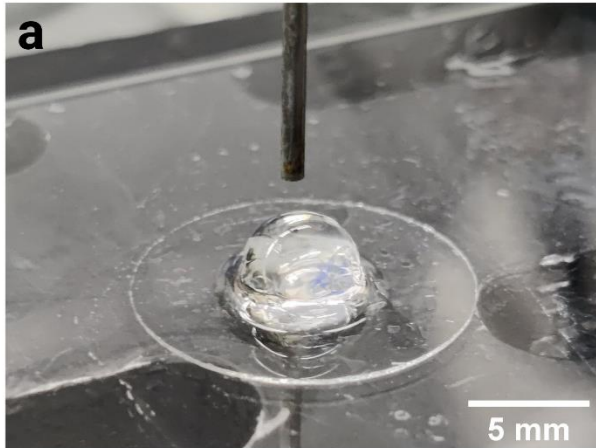
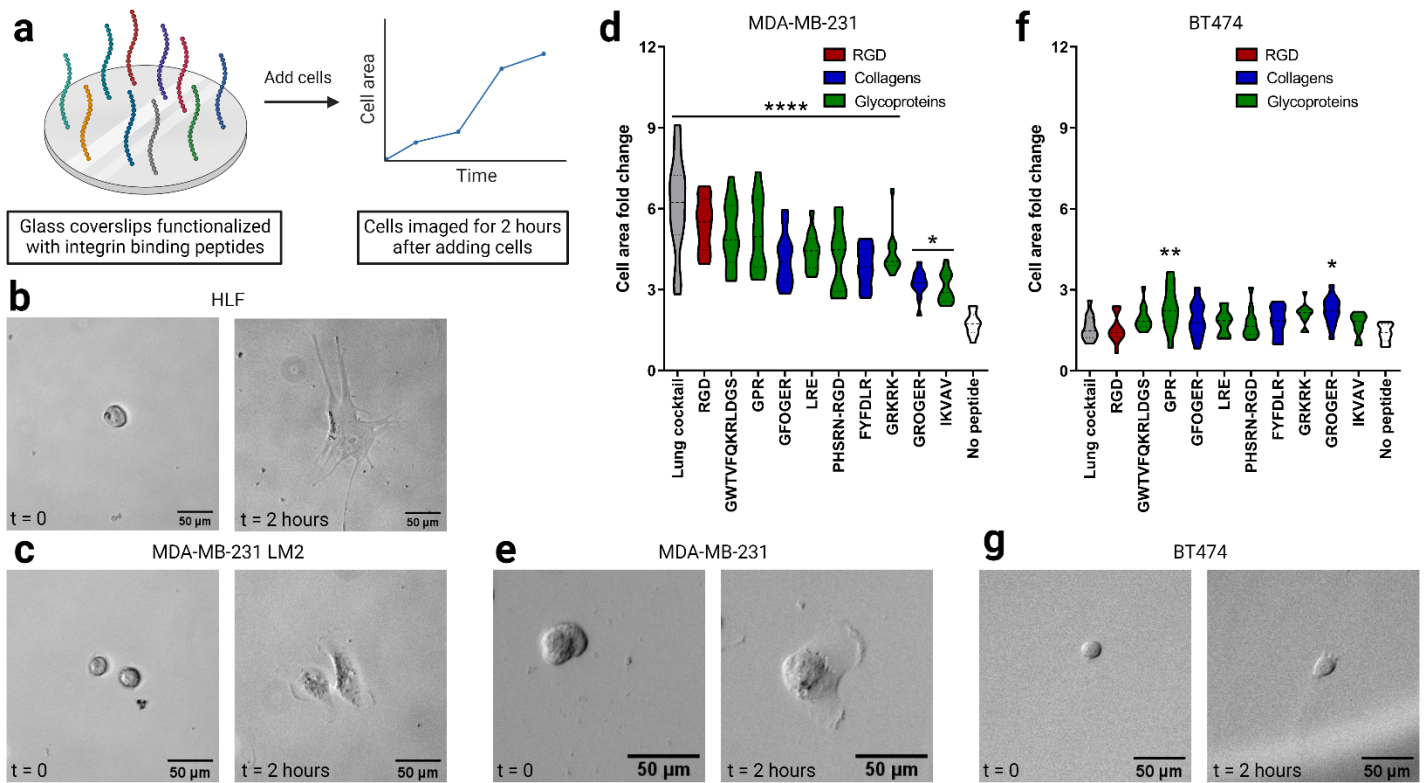


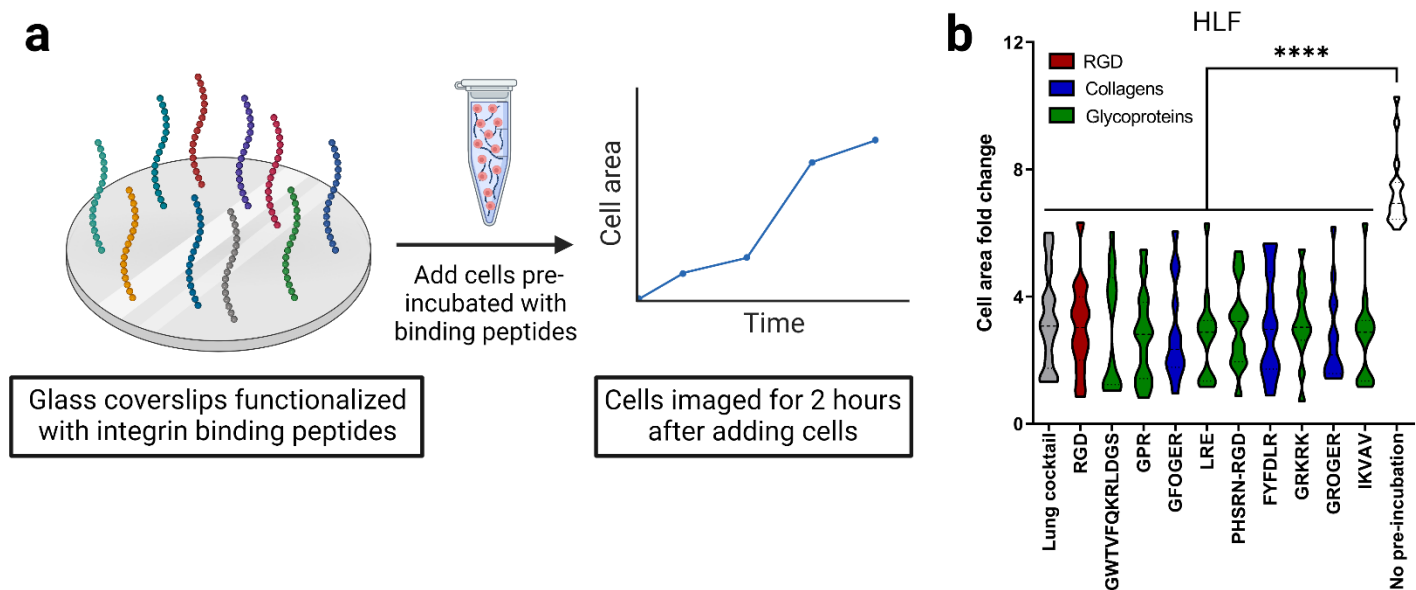
Supporting Information



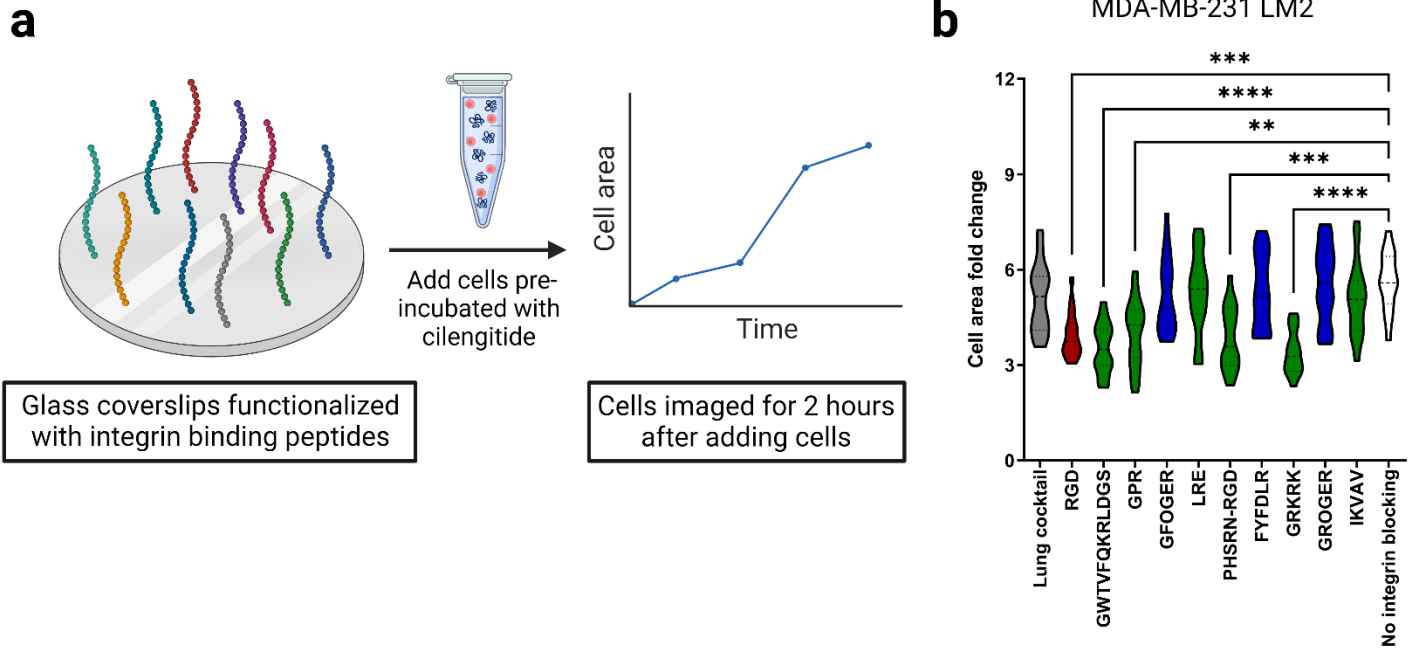
Supplementary Figure 1. Mechanical characterization of lung hydrogels using micro-indentation method. a) Representative image of lung hydrogel after 24 hours of swelling in 1X PBS. Modulus of the hydrogel was measured using micro-indentation method with a 1 mm diameter flat probe. b) Effective modulus of 10 wt% 4-arm PEG-maleimide hydrogel without bioactive peptide ligands using micro-indentation replicated the lung modulus measured using the same method. c) Effective modulus of 10 wt% 4-arm PEG-maleimide hydrogel containing different molar concentrations of MMP-degradable peptide crosslinkers using micro-indentation method. The tested peptide crosslinker molar concentrations were 0%, 13% and 25%. d) Effective modulus of 10 wt% 4-arm PEG-maleimide hydrogel containing different molar concentrations of integrin-binding peptide moieties using micro-indentation method. The tested peptide molar concentrations were 0 mM, 1 mM, 2 mM, 4 mM, and 6 mM.



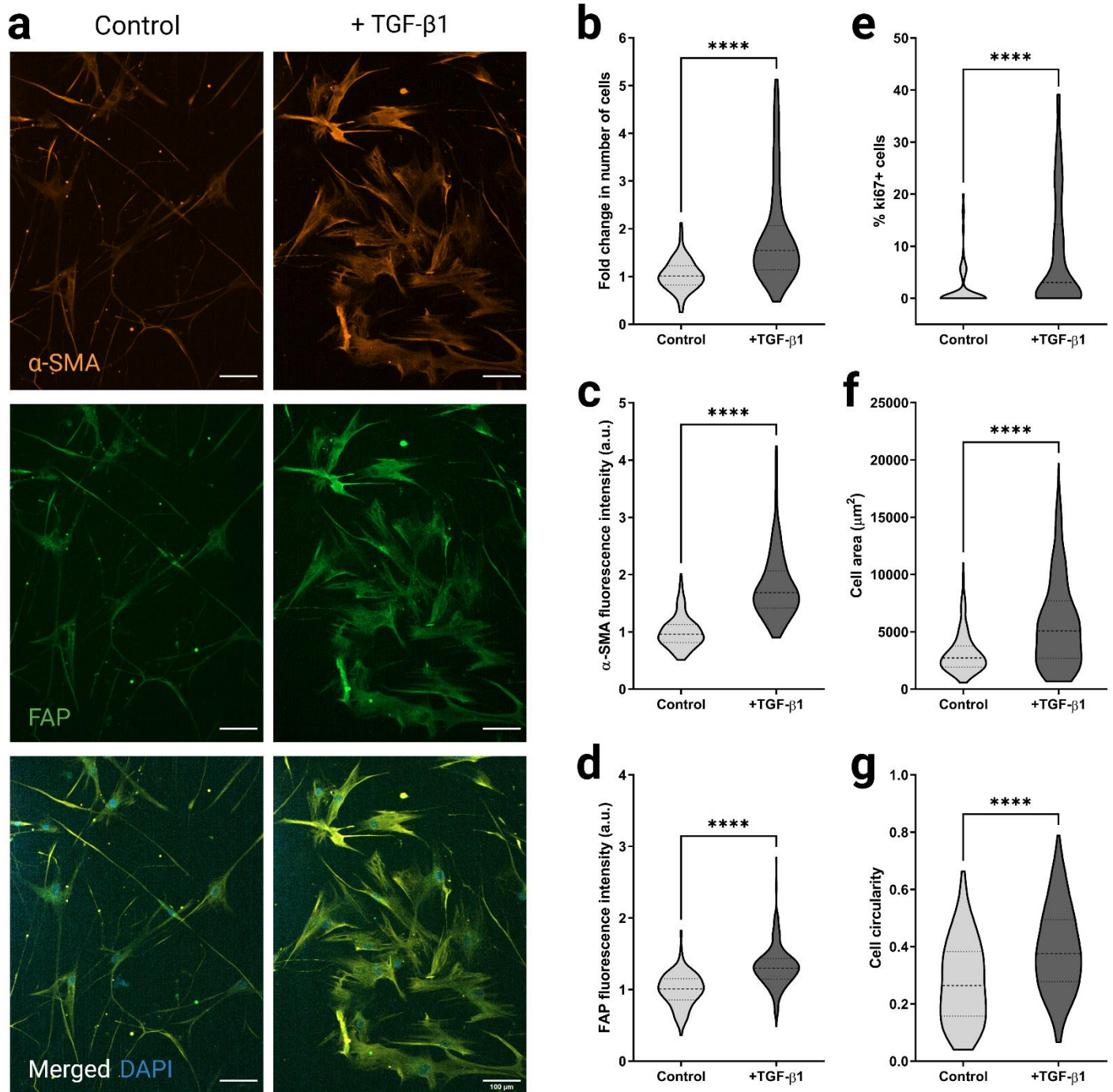
Supplementary Figure 2. Functional validation of lung integrin-binding peptides. a) Schematics of the cell attachment assay where cells were seeded on glass coverslips functionalized with lung integrin-binding peptides and the cells were imaged for 2 hours. b) Representative images of HLFs at time $t = 0$ and $t = 2$ hours. c) Representative images of MDA-MB-231 LM2 cells at time $t = 0$ and $t = 2$ hours. d) Cell area fold change for MDA-MB-231 cells in compared to negative control. e) Representative images of MDA-MB-231 cells at time $t = 0$ and $t = 2$ hours. f) Cell area fold change for BT474 cells in compared to negative control. g) Representative images of BT474 cells at time $t = 0$ and $t = 2$ hours.



Supplementary Figure 3. Competitive cell attachment assay on lung integrin-binding peptides using HLFs. a) Schematics of the competitive cell binding assay where HLFs pre-incubated with individual lung integrin-binding peptides were seeded on glass coverslips functionalized with lung integrin-binding peptide cocktail and the cells were imaged for 2 hours. b) Representative images of HLFs (pre-incubated with individual binding peptides) at time $t = 0$ and $t = 2$ hours.



Supplementary Figure 4. Integrin-specific cell attachment with lung integrin-binding peptides. a) Schematics of the competitive cell binding assay where HLFs pre-incubated with cilengitide were seeded on glass coverslips functionalized with lung integrin-binding peptide cocktail and the cells were imaged for 2 hours. b) Cell area fold change 2 hours after seeding HLFs (pre-incubated with cilengitide) onto glass coverslips functionalized with integrin-binding peptide cocktail relative to a negative control (cells not pre-incubated with cilengitide).



Supplementary Figure 5. Fibroblast phenotype and activation on TCPS. (a) Representative fluorescent images of HLFs cultured on TCPS with or without pro-fibrotic cytokine, TGF- β 1 showing α -SMA (orange), and FAP (green) expressions along with merged HLF images with nuclei staining with DAPI (blue). (b) Fold change in cell count for non-activated (cultured without TGF- β 1) and activated (cultured with TGF- β 1) HLFs on TCPS representing cell proliferation characteristics. (c) Quantification of α -SMA expression from non-activated and activated HLFs cultured on TCPS. (d) Quantification of FAP expression from non-activated and activated HLFs cultured on TCPS. (e) Percentage of proliferative ki67+ cells in non-activated and activated HLF cultures on TCPS. (f) Cell area for non-activated and activated HLFs cultured on TCPS. (g) Cell circularity for non-activated and activated HLFs cultured on TCPS.



b

ECM proteins	with known	integrin-binding domains
Collagen VI ($\alpha 1$, $\alpha 2$, and $\alpha 3$), Emilin I, Fibrillin I, Fibrinogen β , Nidogen I, Nephronectin, Vitronectin, von Willebrand Factor	Fibrinogen α	Collagen IV ($\alpha 1$, $\alpha 2$)
	GPRGGC , 8%	GCGFYFDLR , 4%
	Collagen I ($\alpha 1$, $\alpha 2$)	Elastin
GRGDSPCG , 32%	CGP(GPP) ₅ GFOGER (GPP) ₅ , 6%	GCGRKRK , 2%
Tenascin C	Laminin γ	Collagen III ($\alpha 1$)
CGGAEIDGIEL , 22%	GCKQLREQ , 6%	CGP(GPP) ₅ GROGER (GPP) ₅ , 2%
Fibrinogen γ	Fibronectin	Laminin α
GCGWTVFQKRLDGS , 10%	CGPHSRNG ₅ RGDS , 5%	CSRARKQAASIKVAVADR, 2%

c

Synthesize	peptides	degradable by target	MMPs
MMP1		MMP2	
GCRDVPMS/MRGGDRCG , 16%		GCRDSGESPAY/YTADRRCG , 16%	
MMP3		MMP7	
GCRDRPFS/MIMGDRCG , 16%		GCRDVPLS/LTMGDRCG , 16%	
MMP9		MMP13	
GCRDVPLS/LYSGDRCG , 12%		GCRDGPLG/LWARDRCG , 13%	
		MMP14	
		GCRDIPES/LRAGDRCG , 12%	

Supplementary Figure 6. List and quantities of metastatic lung-specific integrin-binding and MMP-degradable peptides. a) Schematic of identification of metastatic lung ECM-specific integrin-binding and MMP-degradable peptides. b) List of identified integrin-binding peptide domains with their relative quantities corresponding to metastatic lung ECM proteins. c) List of identified MMP-degradable peptide domains with their relative quantities corresponding to the different MMPs.