

Supplementary Materials for
**Injectable hydrogel microspheres with self-renewable hydration layers
alleviate osteoarthritis**

Yiting Lei, Yuping Wang, Jieliang Shen, Zhengwei Cai, Chen Zhao, Hong Chen, Xiaoji Luo,
Ning Hu*, Wenguo Cui*, Wei Huang*

*Corresponding author. Email: huncqjoint@yeah.net (N.H.); wgcui80@hotmail.com (W.C.);
huangw511@163.com (W.H.)

Published 2 February 2022, *Sci. Adv.* **8**, eabl6449 (2022)
DOI: 10.1126/sciadv.abl6449

This PDF file includes:

Figs. S1 to S9
Table S1

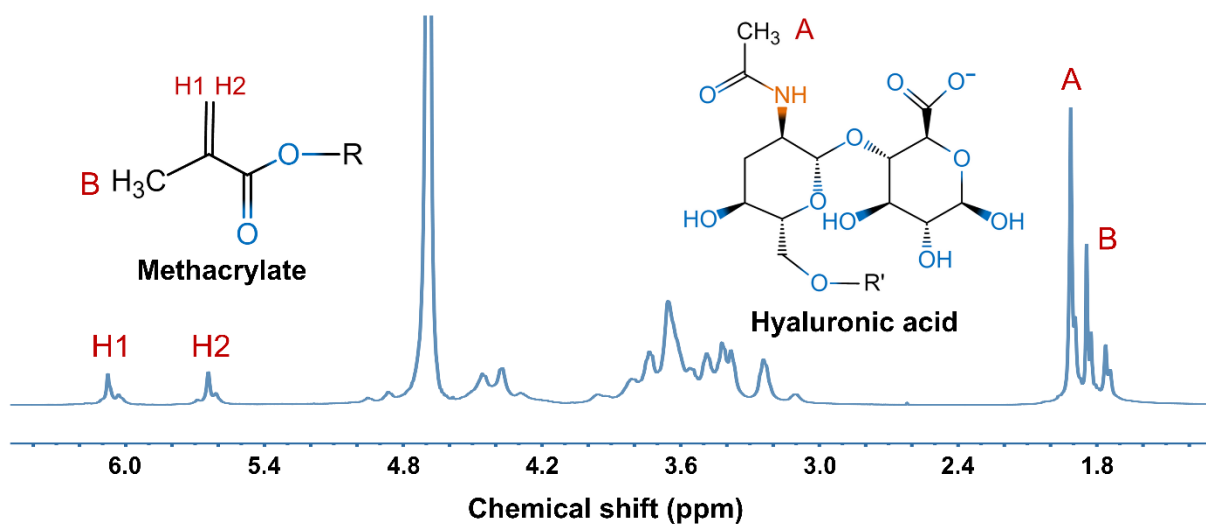


fig. S1. The ¹H NMR spectra of HAMA.

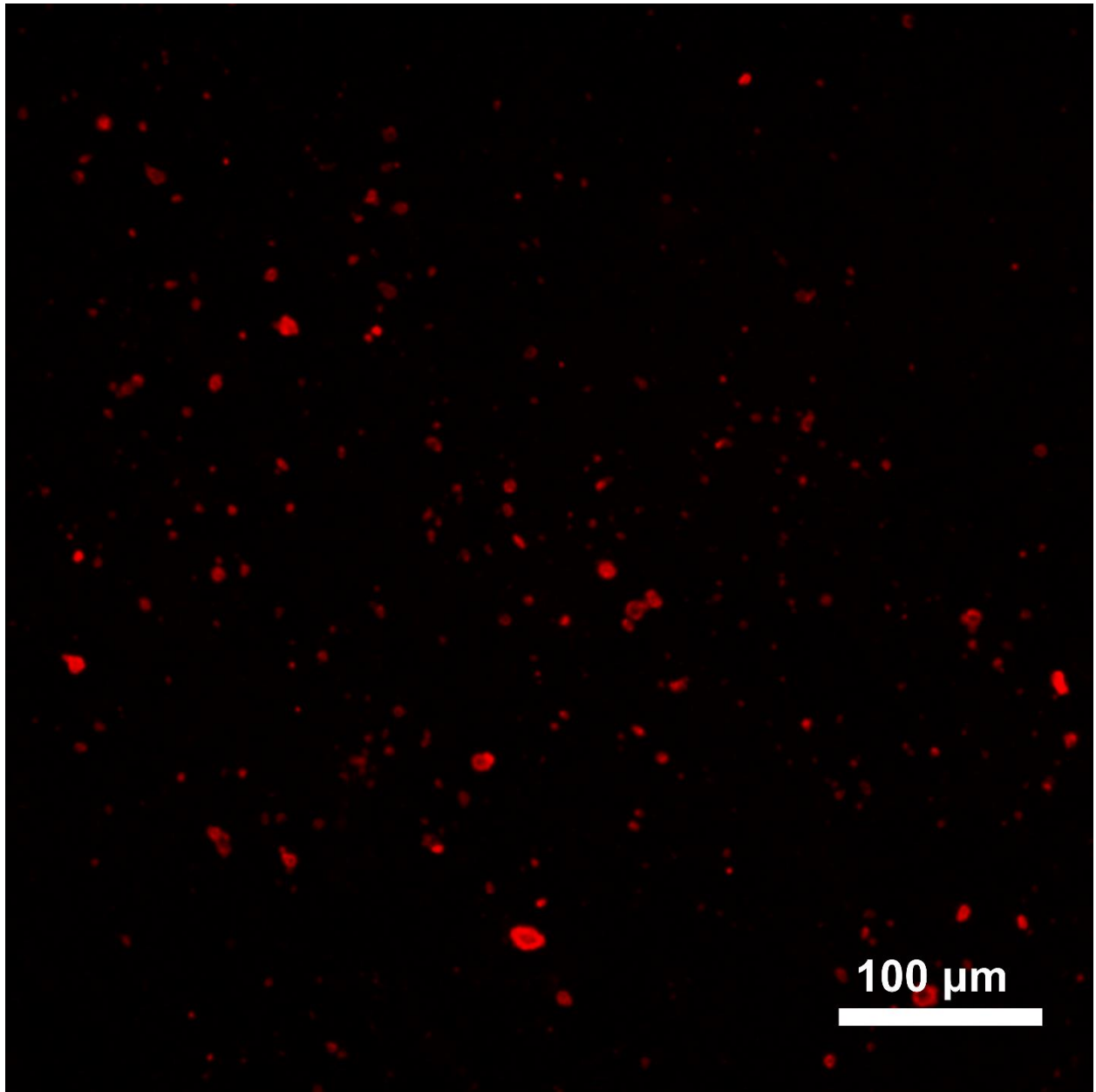


fig. S2. LSCM image of the cartilage section incubated with Dil-labelled cationic liposomes.

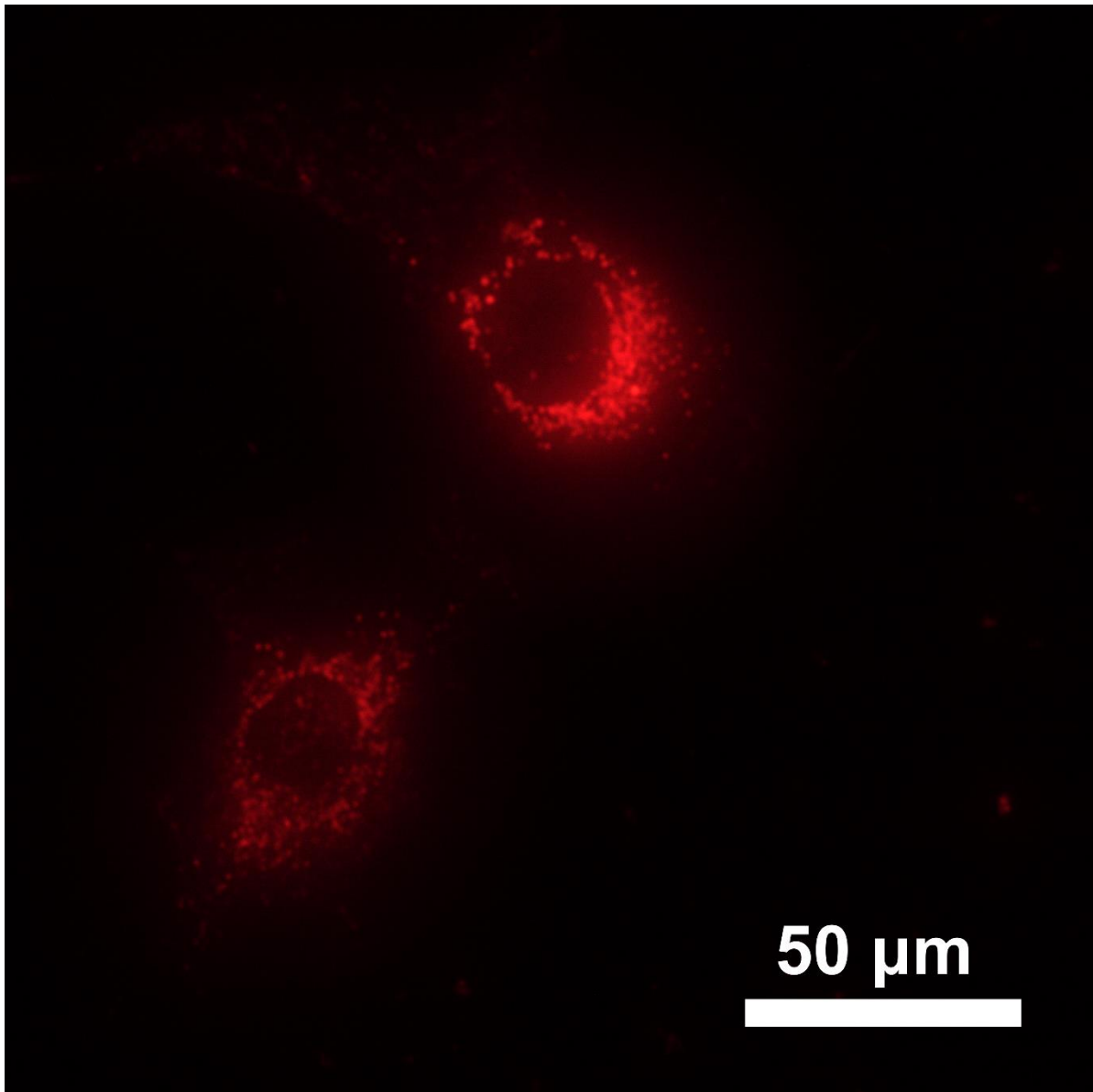


fig. S3. LSCM image of the chondrocytes incubated with Dil-labelled cationic liposomes.

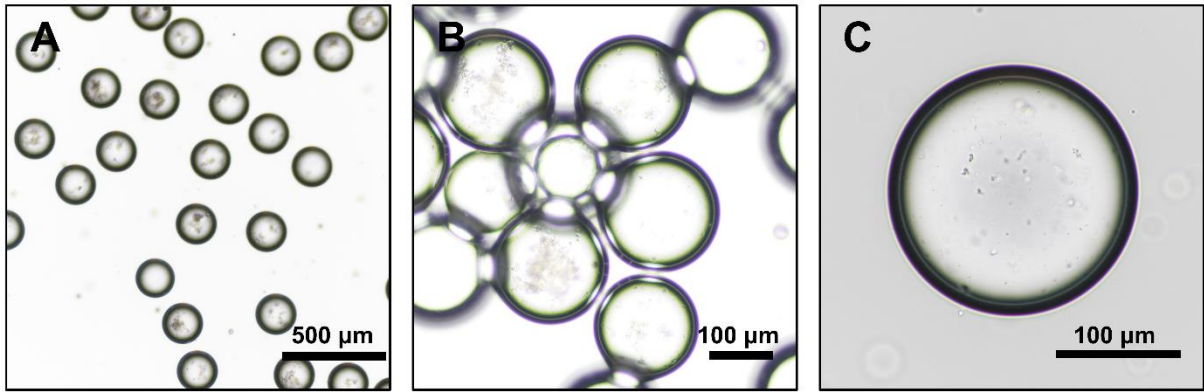


fig. S4. Characterization of pre-gel droplets: (A-B) dispersed pre-gel droplets, and (C) mono-pre-gel droplet.

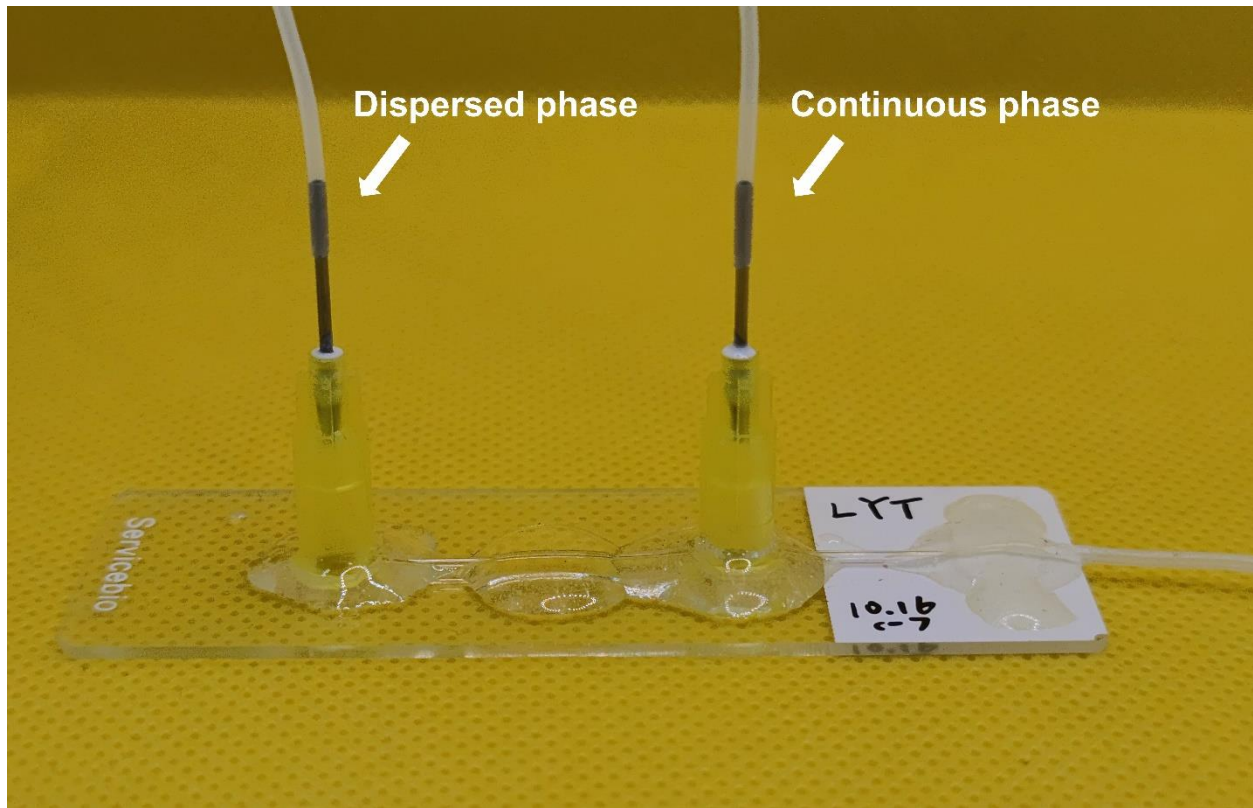


fig. S5. Photograph of the microfluidic device. Photo Credit: Yiting Lei, The First Affiliated Hospital of Chongqing Medical University.

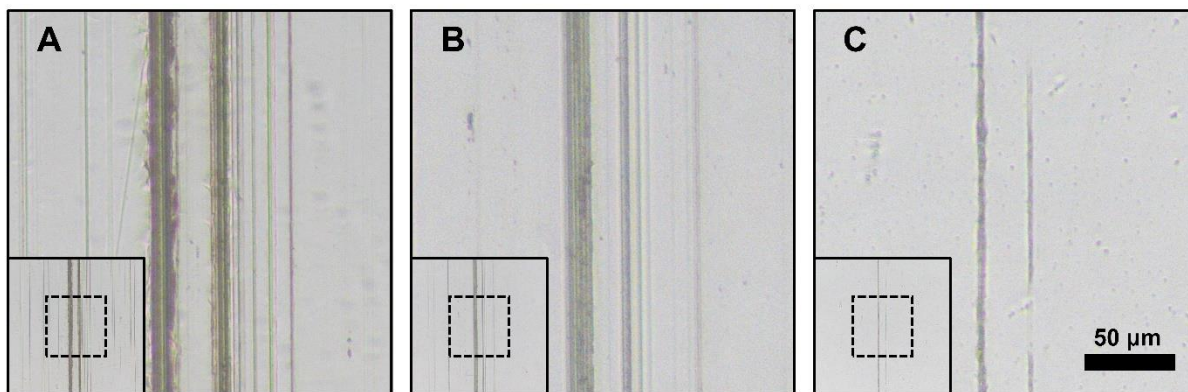


fig. S6. Bright-field images of the wear track on the disk tested by (A) PBS, (B) HMs, and (C) the worn Lipo@HMs.

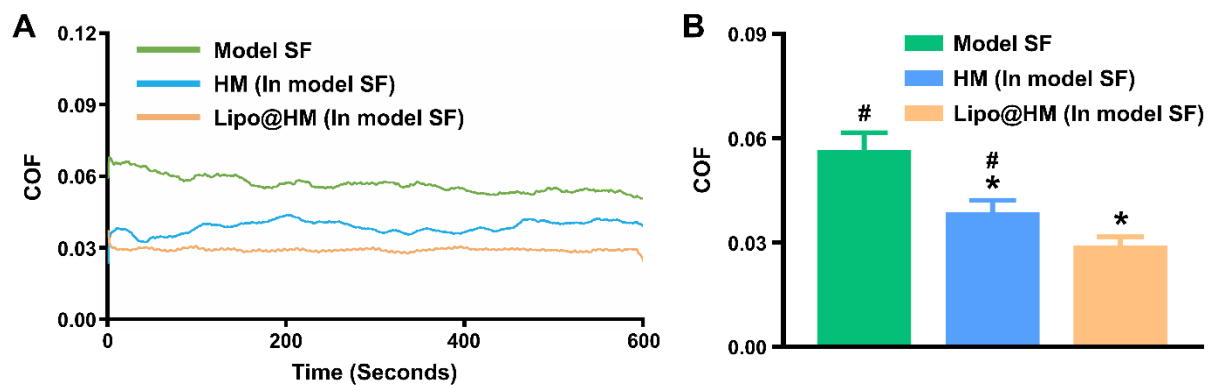


fig. S7. The lubricating performance in the presence of SF: (A) COF-time curves and (B) COF histograms for model SF, HMs (in model SF), and the worn Lipo@HMs (in model SF). (# and * indicated $P < 0.05$ in comparison with the Lipo@HM and SF groups, respectively).

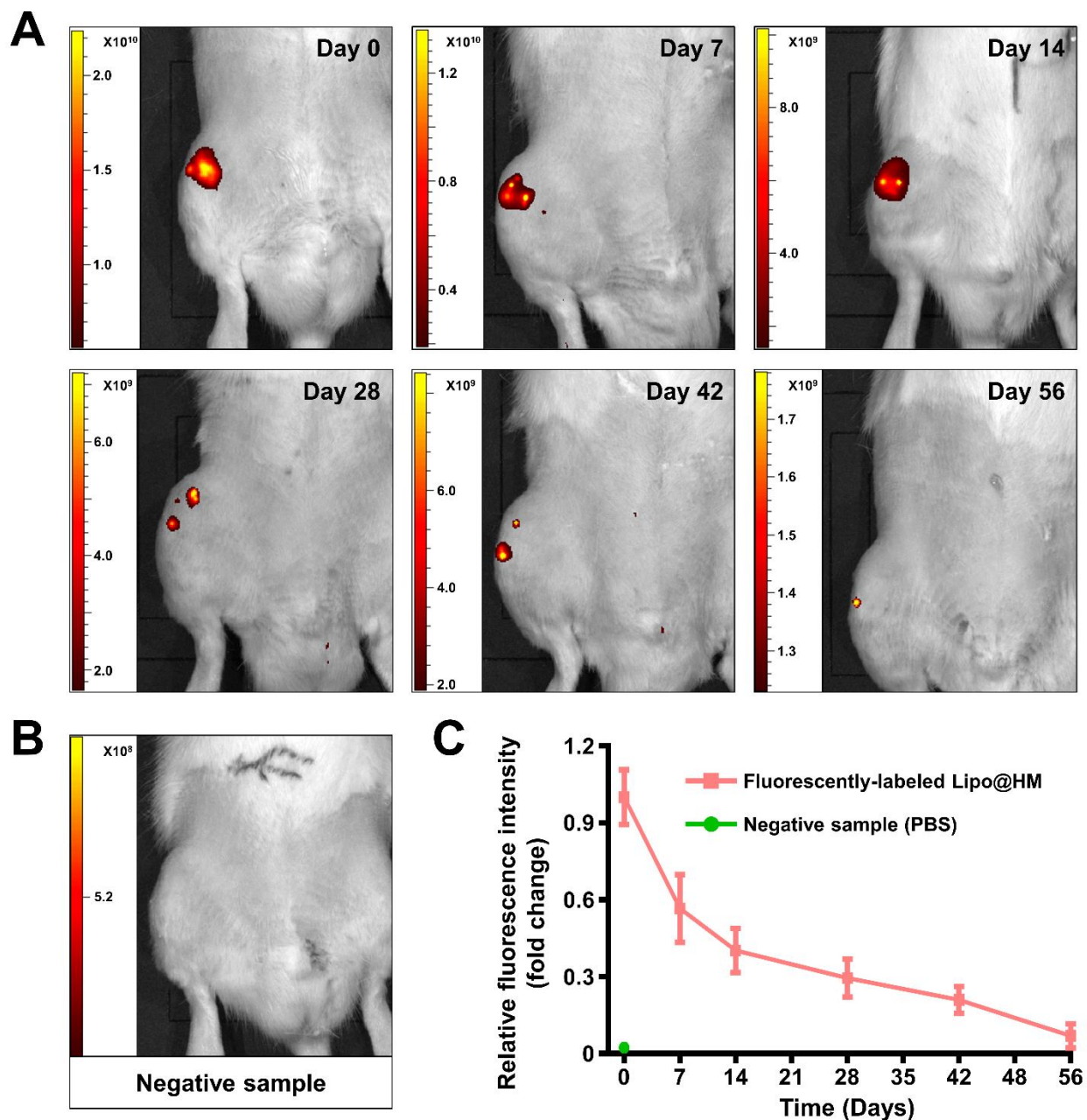


fig. S8. In vivo retention time: (A) The IVIS images of fluorescently labeled Lipo@HMs at different time points. (B) The IVIS image of the negative sample. (C) The relative fluorescence intensity at each time points (relative to fluorescently labeled Lipo@HMs on day 0).

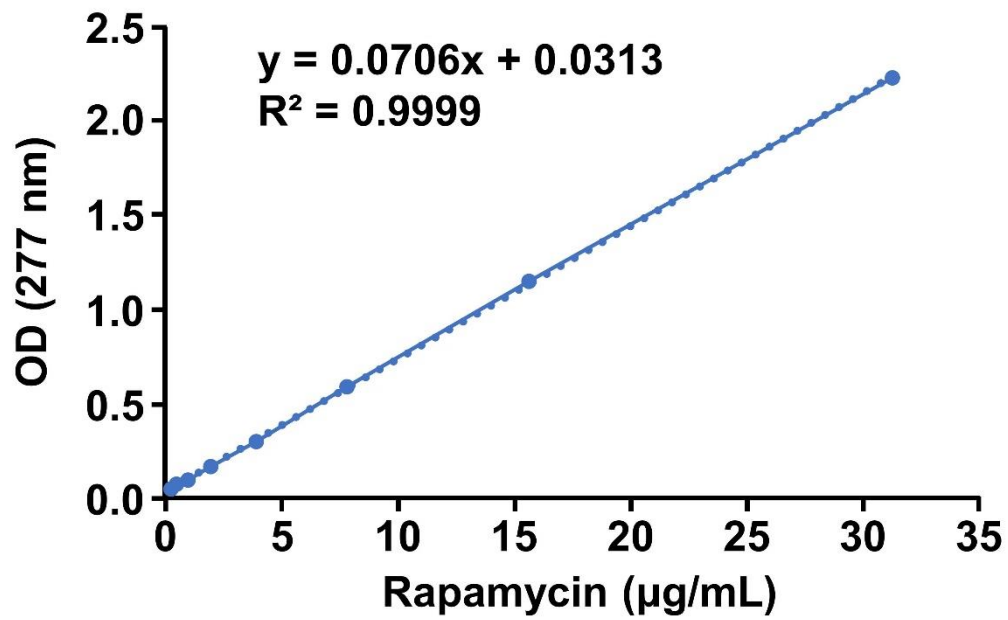


fig. S9. The calibration curve of RAPA with different concentrations.

Table S1. Primers used in real-time PCR.

Gene	Primer	Sequence
GAPDH	Forward	GGAAGCTTGTCATCAATGGAAATC
	Reverse	TGATGACCCTTTTGGCTCCC
Col2	Forward	CACTCAAGTCCCTCAACAACCAG
	Reverse	GGGGTCAATCCAGTAGTCTCCAC
LC3B	Forward	CGAACAAAGAGTAGAAGATGTCCGA
	Reverse	GCTGCTTCTCACCCCTGTATCG
ATG5	Forward	GGATGAGATAACTGAAAGGGAAGC
	Reverse	CCATTTTCAGTGGTGTGCCTTC
MMP13	Forward	GGTGATGAAGATGATTTGTCTGAGG
	Reverse	CGTCAAGTTTGCCAGTCACCT