

Supplementary Information

Cationic micelles as nanocarriers for enhancing intra-cartilage drug penetration and retention

Chenxian Zhu,^a Zhongxing Zhang,^{*a} Yuting Wen,^{a,b} Xia Song,^a Jingling Zhu,^{a,c} Yifei Yao,^{*d} and Jun Li^{*a,b,c}

^a Department of Biomedical Engineering, National University of Singapore, 15 Kent Ridge Crescent, Singapore 119276, Singapore

^b National University of Singapore (Chongqing) Research Institute, 2 Huizhu Road, Yubei District, Chongqing 401120, China

^c NUS Environmental Research Institute (NERI), National University of Singapore, 5A Engineering Drive 1, Singapore 117411, Singapore

^d School of Biomedical Engineering, Shanghai Jiao Tong University, Shanghai 200030, China

***Corresponding authors**

Emails: biezhzh@nus.edu.sg (ZZ), yifeiyao@sjtu.edu.cn (YY), jun-li@nus.edu.sg (JL).

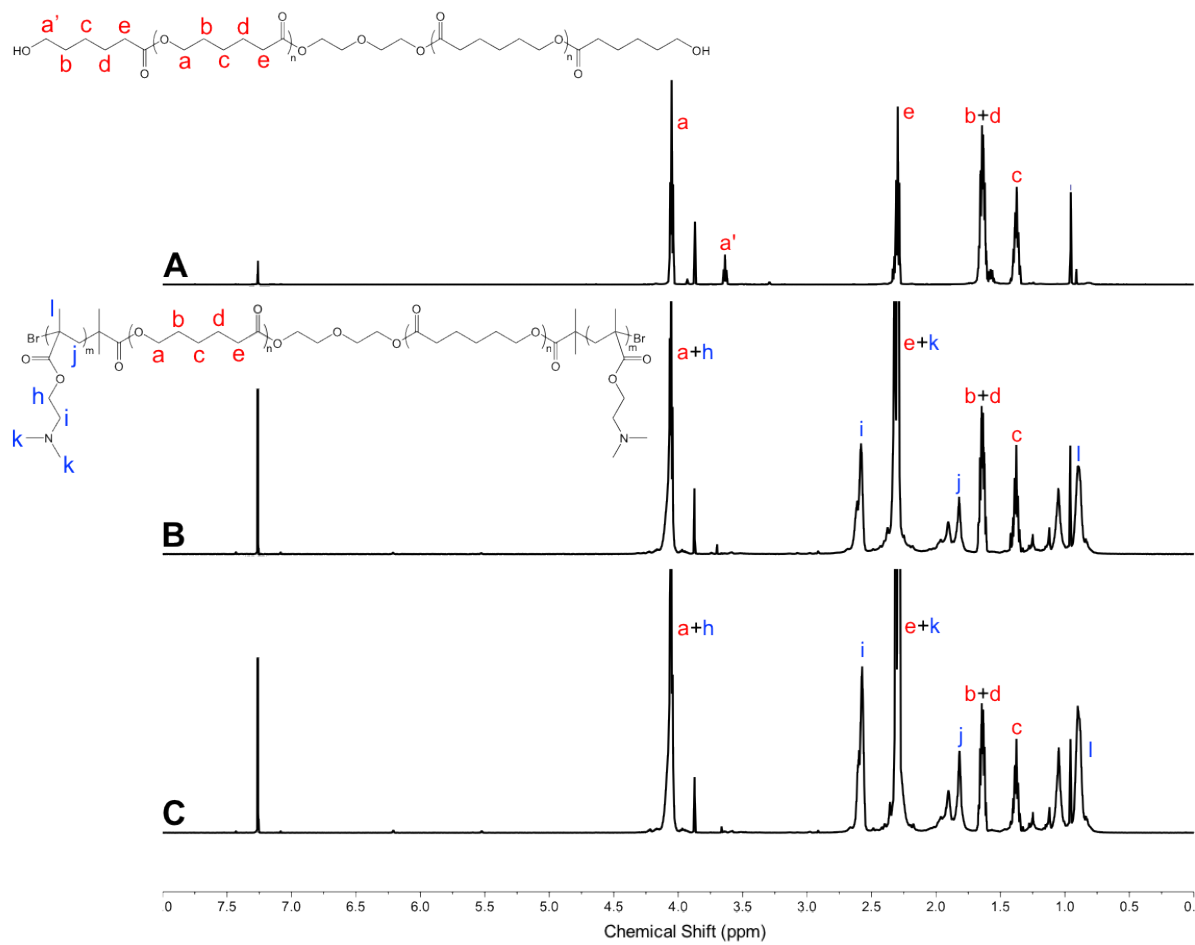


Figure S1. ^1H NMR spectra of polymers in CDCl_3 . (A) PCL-diol; (B) $\text{D}_{16}\text{CL}_{17}\text{D}_{16}$; (C) $\text{D}_{24}\text{CL}_{17}\text{D}_{24}$.

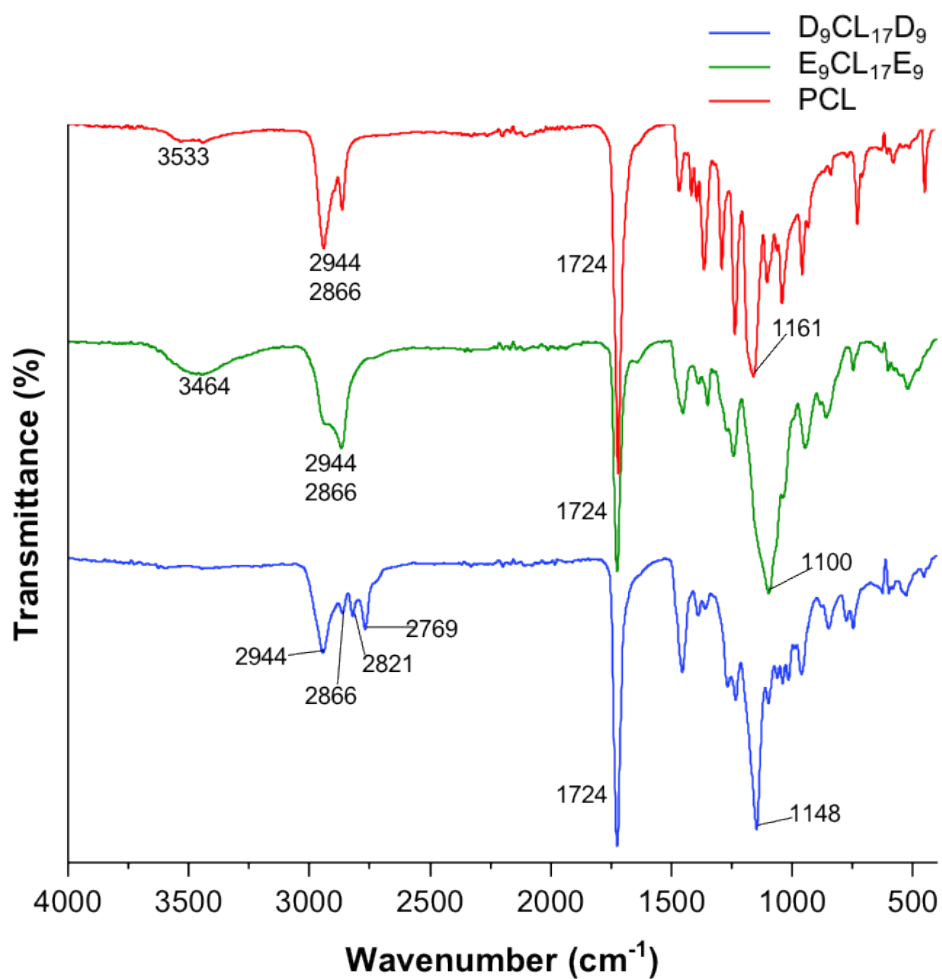


Figure S2. FTIR spectra of PCL-diol, $E_9CL_{17}E_9$, and $D_9CL_{17}D_9$.

Table S1. Zeta potential of $D_9CL_{17}D_9$ and $E_9CL_{17}E_9$ micelles at different pH values. Micelles were prepared by the nanoprecipitation method in $0.1 \times$ PBS at a concentration of 1 mg/mL.

Copolymer	Zeta potential (mV)		
	pH = 5	pH = 7	pH = 9
$D_9CL_{17}D_9$	20.3	17.6	14.1
$E_9CL_{17}E_9$	-0.4	-0.6	-1.4

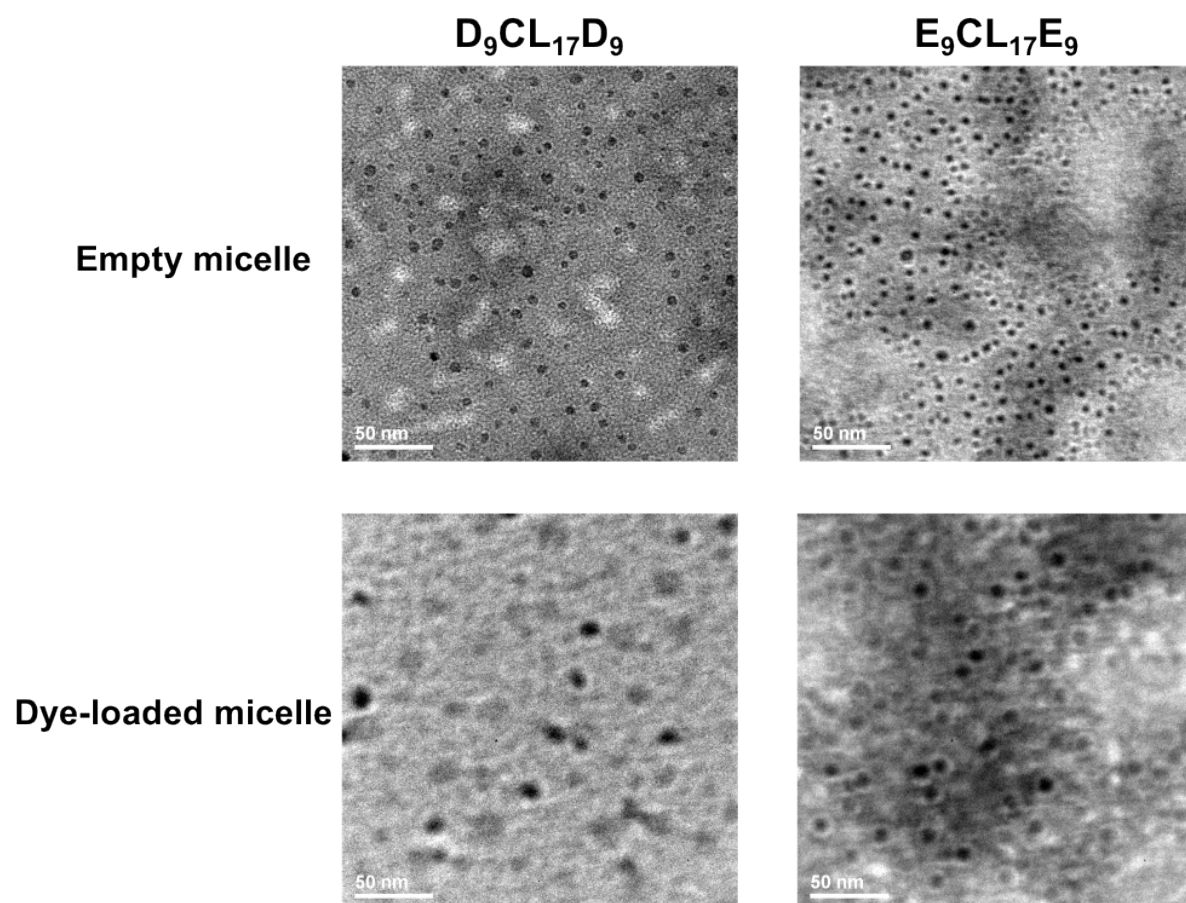


Figure S3. TEM images of empty and dye-loaded $D_9CL_{17}D_9$ and $E_9CL_{17}E_9$ micelles. Scale bar = 50 nm.

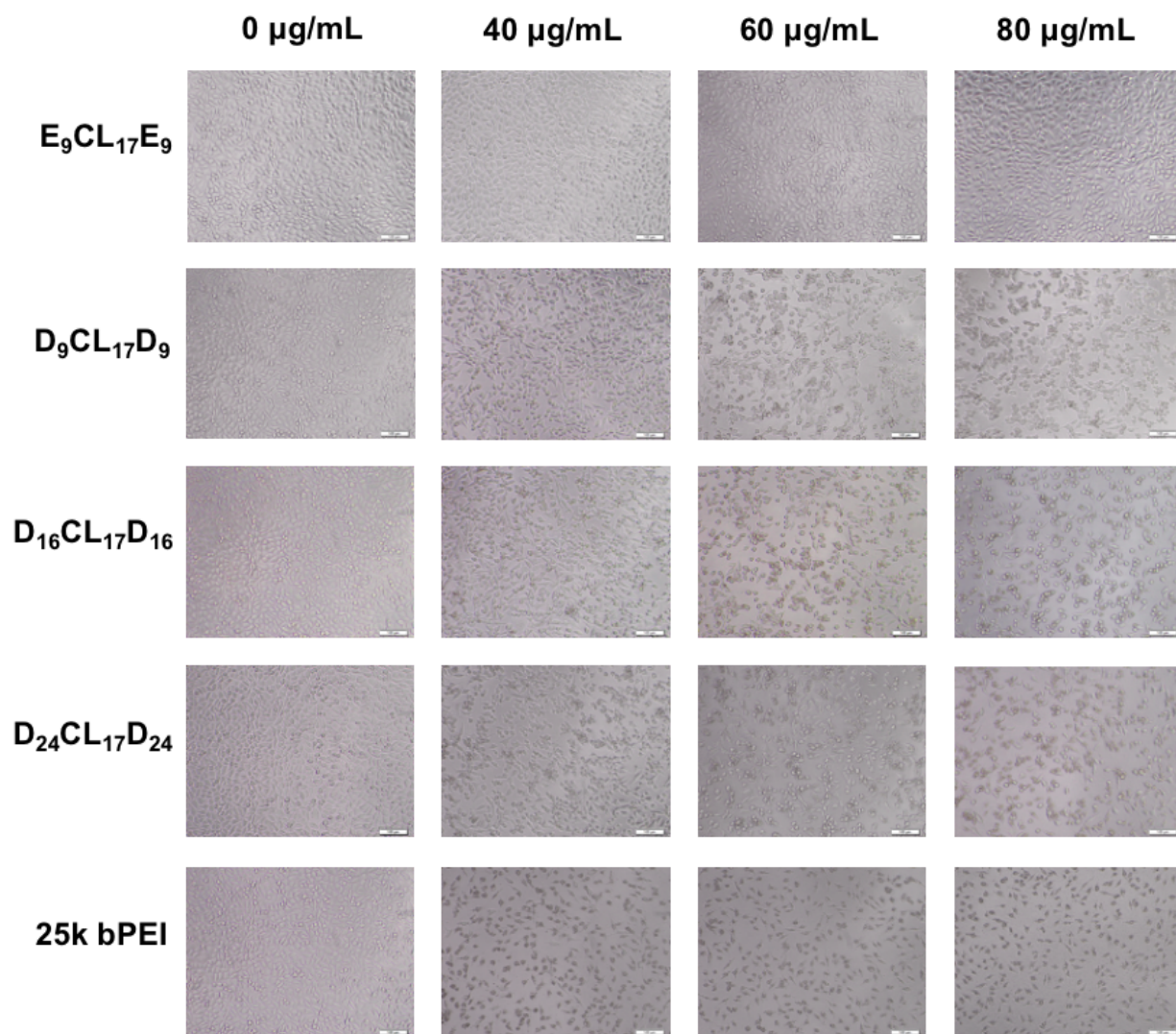


Figure S4. Cell morphology after treated with different micelles and bPEI was observed by microscopy at bright field at $10 \times$ magnification. Scale bar = $100 \mu\text{m}$.

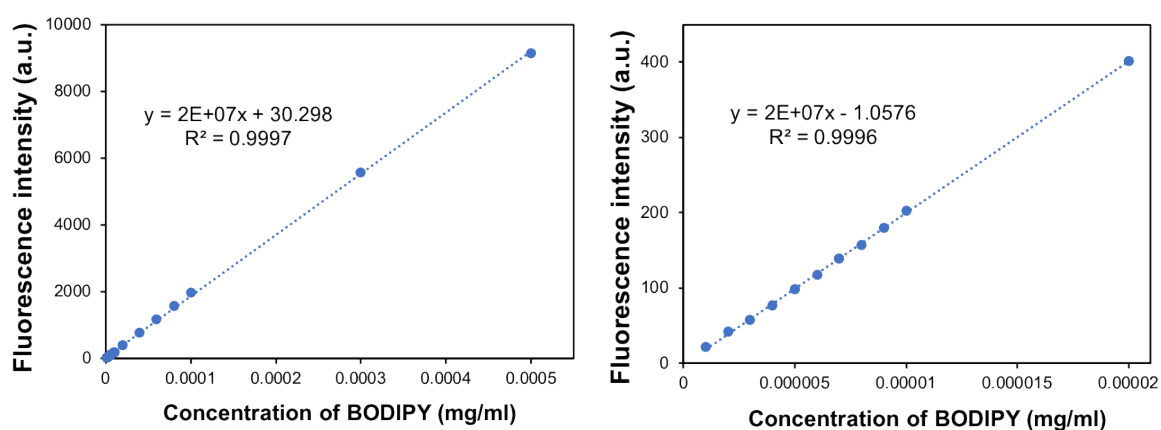


Figure S5. The standard curve of BODIPY with two concentration ranges in a mixture of 20% v/v DI water and 80% v/v acetone.

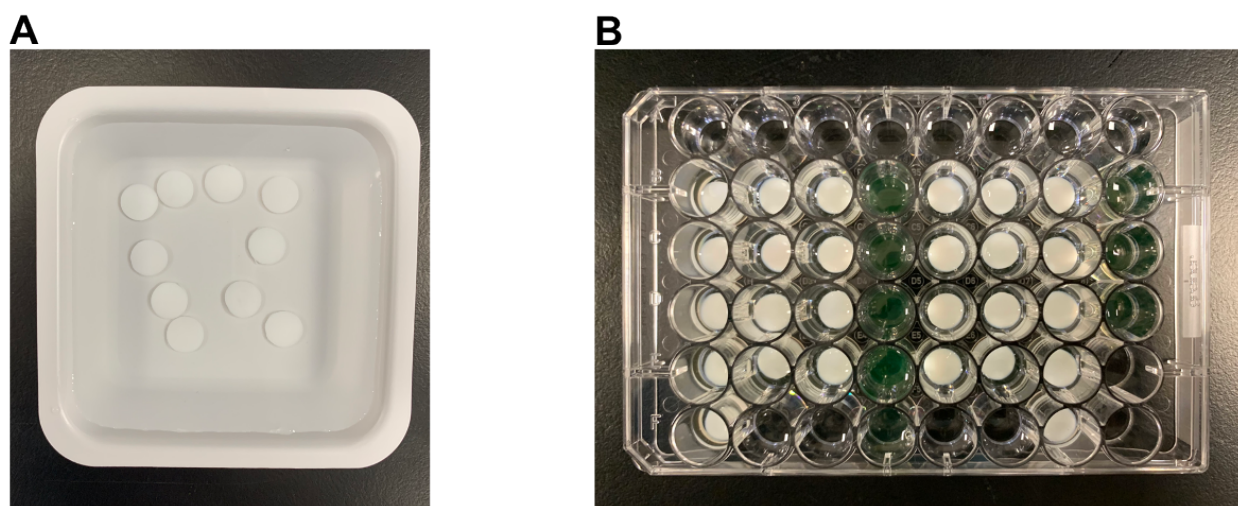


Figure S6. Photos of excised porcine cartilage disks. (A) After harvesting, the disks ($\varnothing 9 \times 1.5$ mm) were washed and equilibrated with a sterile PBS solution. (B) For the absorption or desorption experiments, cartilage disks were equilibrated in 300 μ L of absorption or desorption baths in the 48-well plate.