Supporting Information

Acid Degradable Poly(vinylcaprolactam)-based Nanogel with Ketal Linkage for Drug Delivery

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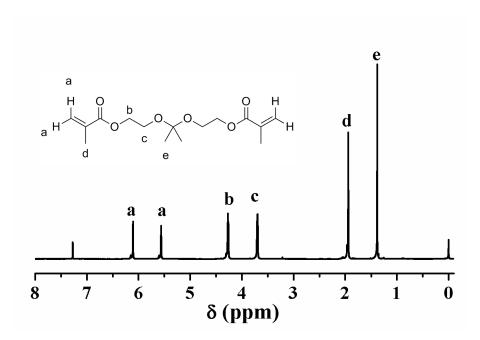


Fig. S1 ¹H NMR spectrum of DMAEP. ¹H NMR (500 MHz, CDCl3): δ (ppm) 1.38 (6H, s, (COOCH₂CH₂O)₂C(CH₃)₂), 1.94 (6H, s, (OCOCCH₃CH₂)₂), 3.70 (4H, m, (COOCH₂CH₂O)₂C(CH₃)₂), 4.27 (4H, m, (COOCH₂CH₂O)₂C(CH₃)₂), 5.57 (2H, s, (OCOCCH₃CH₂)₂ – syn to methyl), 6.11 (2H, s, (OCOCCH₃CH₂)₂ – anti to methyl).

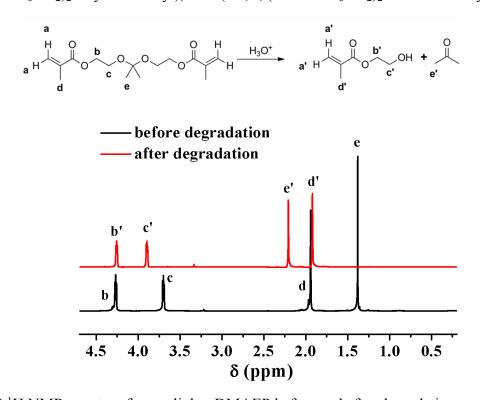


Fig. S2 ¹H NMR spectra of cross-linker DMAEP before and after degradation.

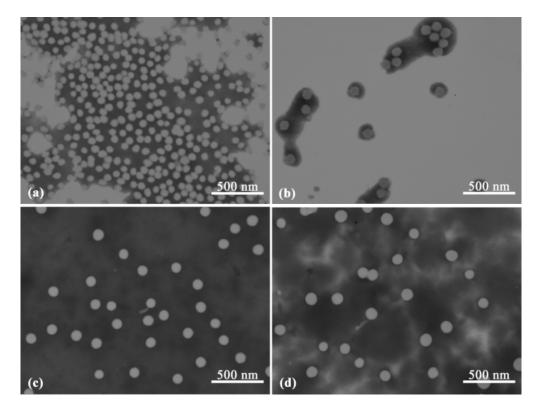


Fig. S3 Representative TEM images (×5W) of P(VCL-*ketal*-HPMA) nanogels with different HPMA contents: (a) 3 wt%, (b) 5 wt%, (c) 8 wt% and (d) 10 wt%.

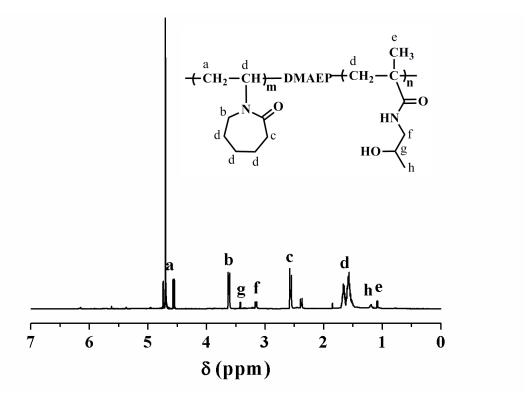


Fig. S4 ¹H NMR spectrum of P(VCL-ketal-HPMA) nanogel in D₂O.

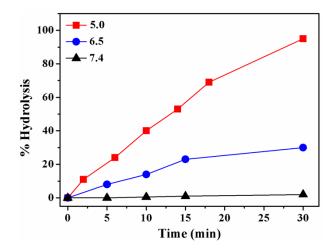


Fig. S5 Hydrolysis kinetics of acid-cleavable cross-linker DMAEP at pH 5.0, 6.5 and 7.4, respectively.

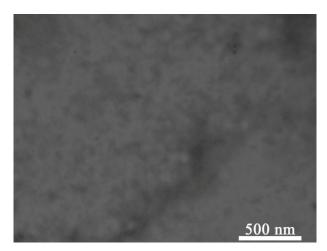


Fig. S6 TEM image of P(VCL-ketal-HPMA-8) nanogels after degradation.

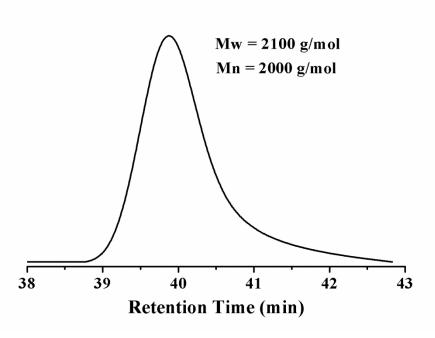


Fig. S7 The molecular weight of copolymer P(VCL-co-HPMA) by GPC measurement.

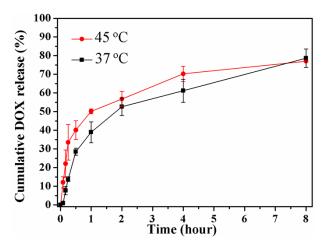


Fig. S8 DOX release profiles of acid-degradable P(VCL-ketal-HPMA) nanogel in the medium of pH 5.0 at 37 °C and 45 °C .

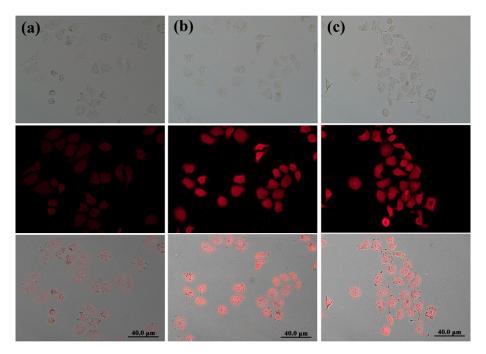


Fig. S9 CLSM images of the intracellular DOX release of free DOX. (a) 2 h, (b) 6 h and (c) 12 h incubation of free DOX. In each column, images from top to down: differential interference contrast microscopy, fluorescence microscopy, and overlays of both images.

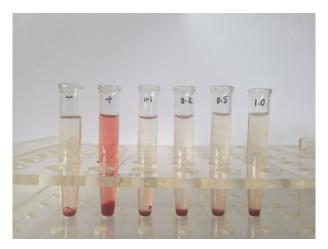


Fig. S10 Hemolysis assay of the P(VCL-*ketal*-HPMA-8) nanogel. RBC suspension to make the final particle concentration 0.1, 0.2, 0.5 and 1 mg/mL, respectively. Distilled water served as positive control and normal saline as negative control.