



## Polymer Chemistry

Electronic Supporting Information for

# Nanoparticles of poly([N-(2-hydroxypropyl)]methacrylamide)-*b*-poly[2-(diisopropylamino)ethyl methacrylate] diblock copolymer for pH-triggered release of paclitaxel

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## Characterization of PHPMA

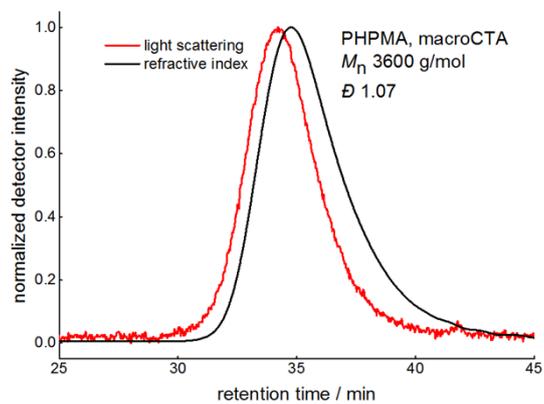


Figure S1A. SEC traces of the PHPMA macroCTA chain transfer agent utilized as macromolecular chain transfer agent for block copolymer synthesis *via* RAFT.

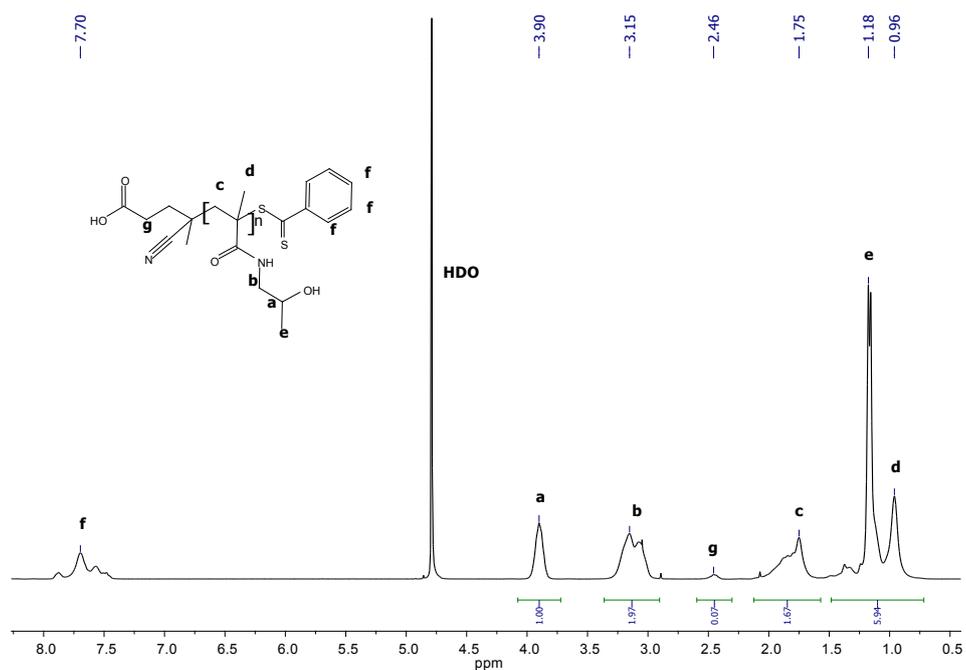


Figure S1B.  $^1\text{H}$  NMR spectra of the PHPMA in  $\text{D}_2\text{O}$ .

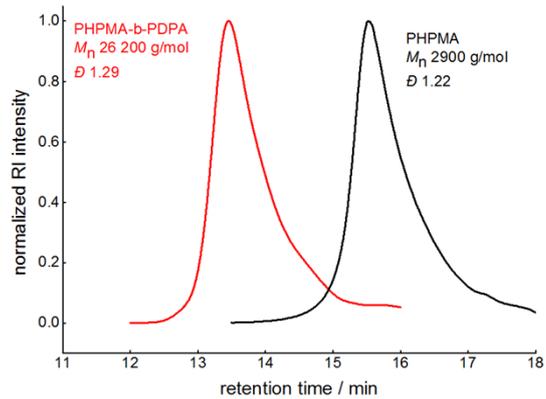


Figure S2A. SEC traces of the PHPMA<sub>25</sub>-*b*-PDPA<sub>106</sub> block copolymer synthesized *via* RAFT.

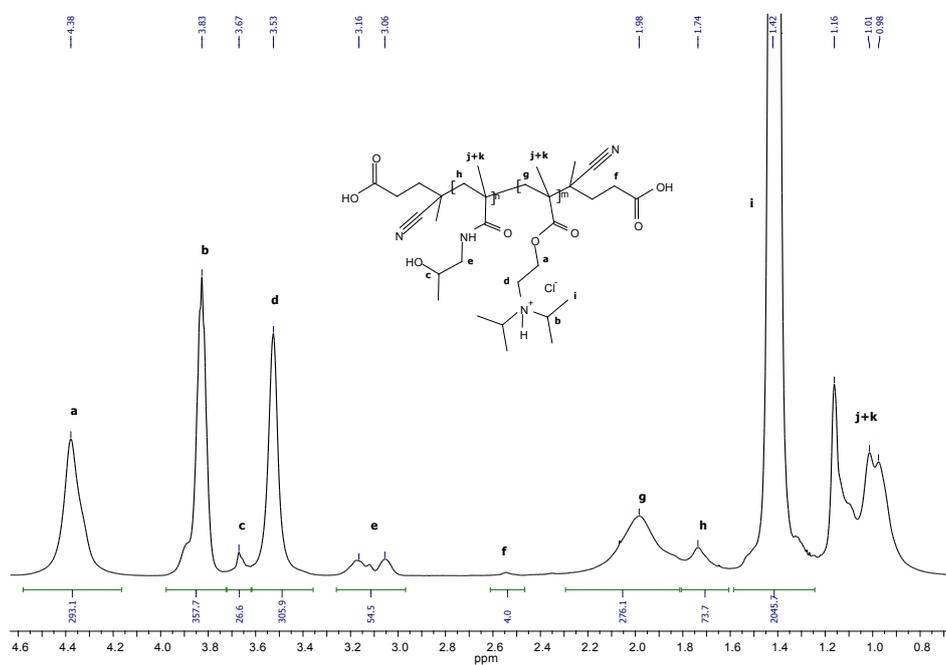


Fig. S2B. <sup>1</sup>H NMR spectra of the PHPMA<sub>25</sub>-*b*-PDPA<sub>106</sub> in D<sub>2</sub>O/DCl (pH 2).

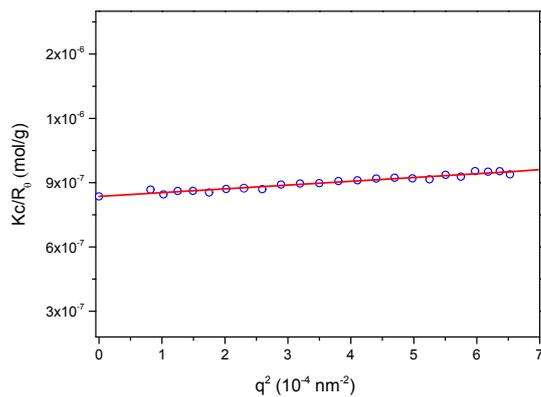


Fig. S3. Static light scattering measurements ( $Kc/R_{90}$  vs.  $q^2$ ) for PHPMA<sub>25</sub>-*b*-PDPA<sub>106</sub> block copolymer NPs in PBS at 25 °C ( $R_G \sim 24$  nm and  $M_w \sim 1.2 \times 10^6$  g/mol).

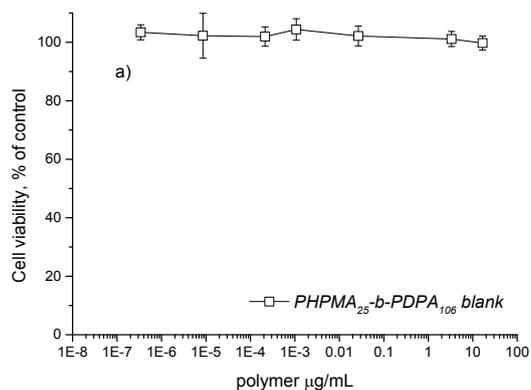


Fig. S4A. Cell viability of HeLa cell line after 24 h incubation with different concentrations of drug-free PHPMA<sub>25</sub>-*b*-PDPA<sub>106</sub> block copolymer NPs.

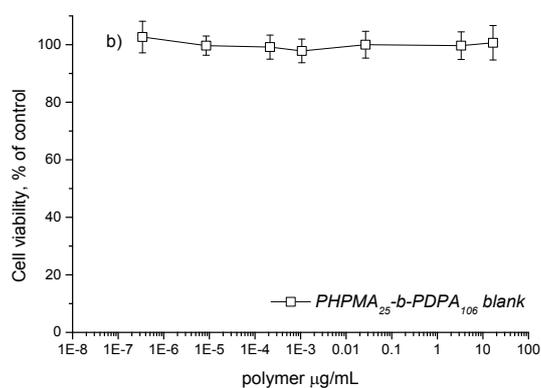
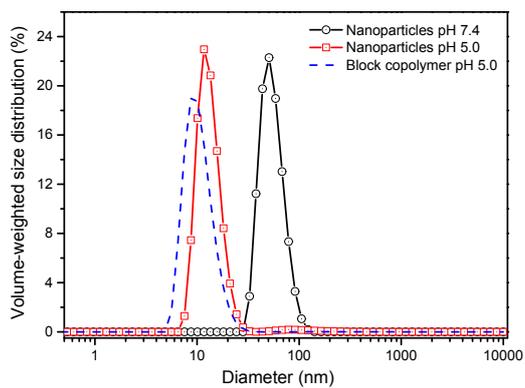
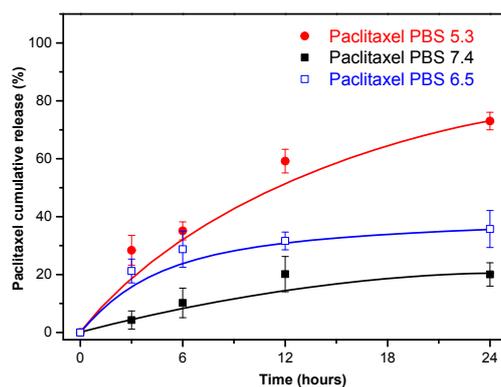


Fig. S4B. Cell viability of HeLa cell line after 48 h incubation with different concentrations of drug-free PHPMA<sub>25</sub>-*b*-PDPA<sub>106</sub> block copolymer NPs.



**Fig. S5.** Volume-weighted size distribution for the PHPMA<sub>25</sub>-*b*-PDPA<sub>106</sub> NPs at pH 7.4 (black open circles) for the nanoparticles at pH 5.0 (red open squares) and the single block copolymer at pH 5.0 (blue dashed lines) and angle 173° at concentration of 1 mg·mL<sup>-1</sup> diluted in PBS at 37 °C.



**Fig. S6.** Drug release profiles from paclitaxel-loaded PHPMA<sub>25</sub>-*b*-PDPA<sub>106</sub> block copolymer NPs at pH of simulated transport in blood, at pH 6.5 (end stage of protonated process) and simulating the acidic environment in endosomal and lysosomal compartments at 37 °C.

**Supporting Table ST1.** Synthetic parameters and molecular weight data of polymers prepared via RAFT polymerization.

Sample	$[M]_0/[CTA]_0/[I]_0$	Time (h)	Conv. (%) <sup>a</sup>	$M_{n,th}$ <sup>b</sup>	$M_{n,SEC}$ <sup>c</sup>	$D$ <sup>c</sup>
				(g mol <sup>-1</sup> )		
PHPMA <sub>25</sub>	120/2/1 <sup>e</sup>	10	36	3 200	3 600 <sup>d</sup> (2 900)	1.07 <sup>d</sup> (1.22)
PHPMA <sub>25</sub> - <i>b</i> -PDPA <sub>106</sub>	300/3/1 <sup>f</sup>	15	94	23 500	26 200	1.29

<sup>a</sup> Determined by <sup>1</sup>H NMR in D<sub>2</sub>O.

<sup>b</sup> Theoretical  $M_n = [M]_0/[CTA]_0 \times \text{conv.} \times MW_{\text{mon.}} + MW_{\text{CTA}}$

<sup>c</sup> Determined by SEC in THF/MeOH 80/20% using PMMA as standard

<sup>d</sup> Determined by SEC in acetate buffer pH 6.5 using light scattering and RI detectors

<sup>e</sup> conditions: DMAc,  $[M]_0 = 1.2$  M, 70 °C

<sup>f</sup> conditions: 1,4-dioxane/MeOH 60/40 vol.%,  $[M]_0 = 3$  M, 70 °C