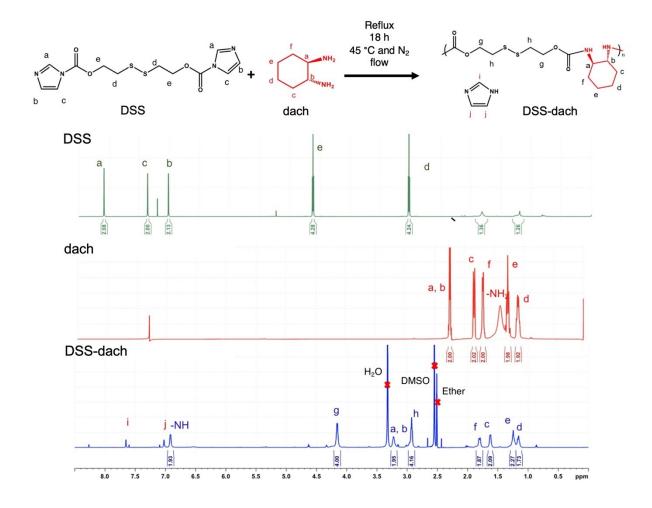
## **Electronic Supporting Information**

## Responsive polyprodrug for anticancer nanocarriers

Arjaree Jobdeedamrong,<sup>a</sup> Man Theerasilp,<sup>a</sup> Nutthanit Thumrongsiri,<sup>b</sup> Paweena Dana,<sup>b</sup> Nattika Saengkrit,<sup>b</sup> Daniel Crespy<sup>a,\*</sup>



**Fig. S1.** <sup>1</sup>H NMR spectrum of DSS-dach in DMSO-d<sub>6</sub>.

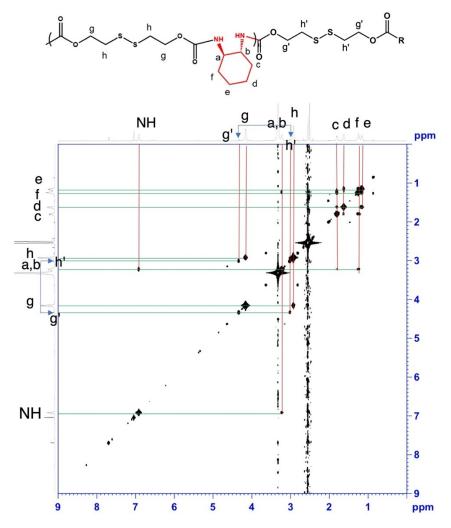


Fig. S2. 2D NMR-COSY spectrum of DSS-dach in DMSO-d<sub>6</sub>.

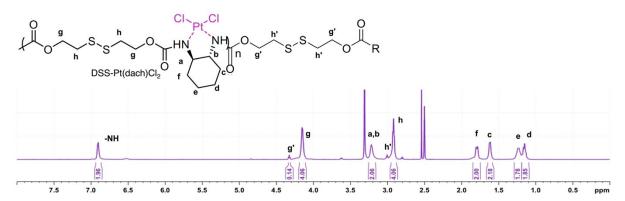
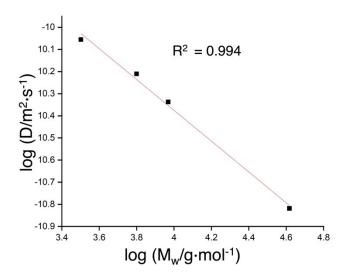
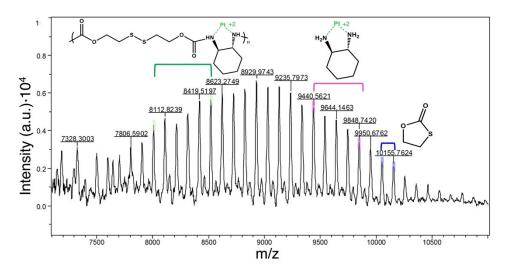


Fig. S3. <sup>1</sup>H NMR spectrum of DSS-Pt(dach)Cl<sub>2</sub> in DMSO-d<sub>6</sub>.



**Fig. S4.** Diffusion coefficients (D) versus molecular weights of poly(ethylene glycol) determined by DOSY-NMR spectroscopy in DMSO-d<sub>6</sub>.



**Fig. S5.** MALDI-TOF mass spectrum of DSS-Pt(dach)Cl<sub>2</sub>. The m/z values represent the mass-to-charge ratios.

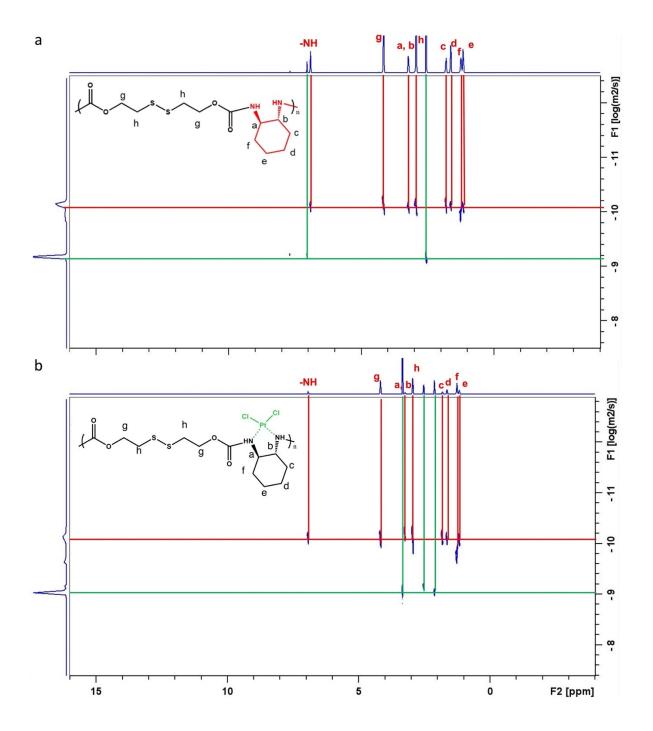


Fig. S6. 2D NMR-DOSY spectra of (a) DSS-dach and (b) DSS-Pt(dach)Cl<sub>2</sub> in DMSO-d<sub>6</sub>.

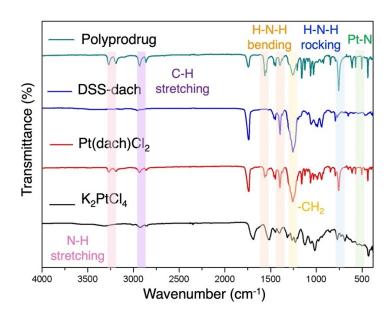


Fig. S7. FT-IR spectra of polyprodrug, DSS-dach, Pt(dach)Cl<sub>2</sub>, and K<sub>2</sub>PtCl<sub>4</sub>.

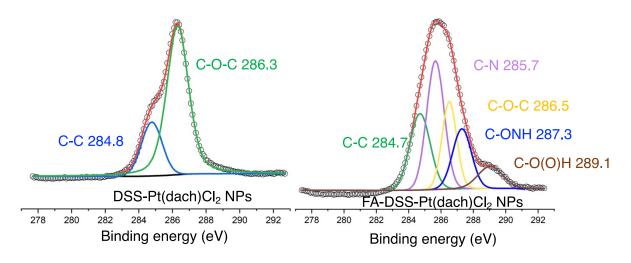


Fig. S8. C 1s XPS spectra of a) DSS-Pt(dach)Cl<sub>2</sub> NPs and b) FA-DSS-Pt(dach)Cl<sub>2</sub> NPs.

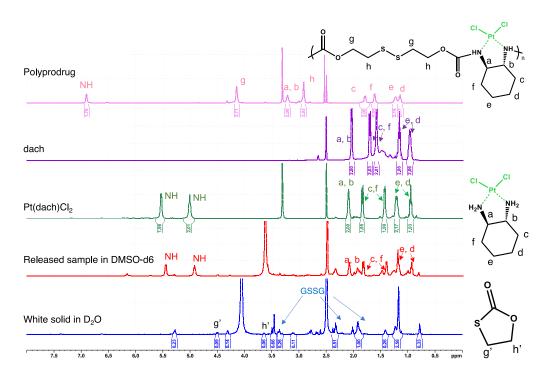
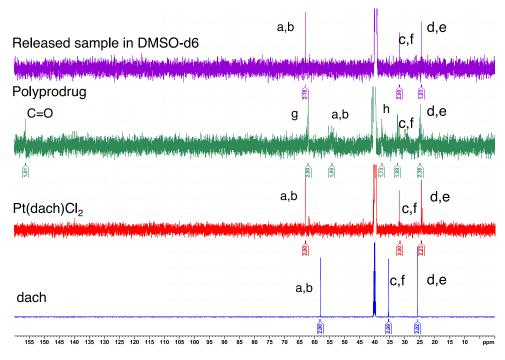


Fig. S9.  $^{1}$ H NMR spectra of polyprodrug, dach, Pt(dach)Cl<sub>2</sub>, and dried release medium after 72 h release in deuterated dimethyl sulfoxide (DMSO-d<sub>6</sub>) and in deuterated water (D<sub>2</sub>O).



**Fig. S10.** <sup>13</sup>C NMR spectra of the dried release medium after release for 72 h, polyprodrug, Pt(dach)Cl<sub>2</sub>, and dach in deuterated dimethyl sulfoxide (DMSO-d<sub>6</sub>).

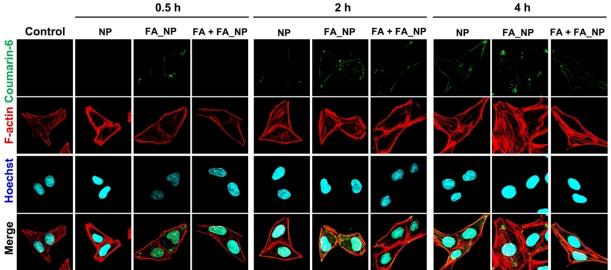
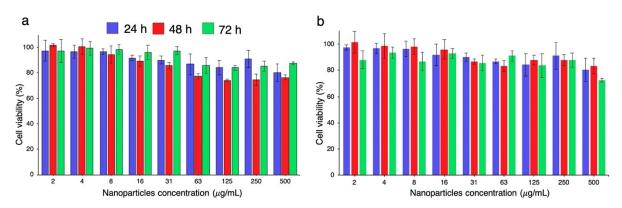
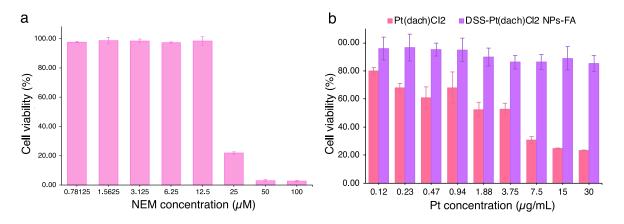


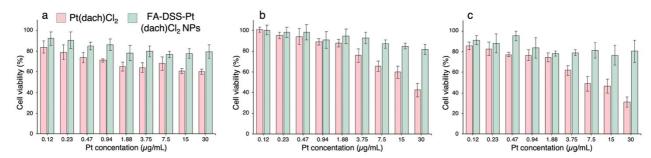
Fig. S11. Cellular uptake images of HeLa cells after 0.5, 2, and 4 h of incubation with DSS-Pt(dach)Cl<sub>2</sub> NPs (NP), FA-DSS-Pt(dach)Cl<sub>2</sub> NPs (FA\_NP), and FA-DSS-Pt(dach)Cl<sub>2</sub> NPs+FA (FA + FA\_NP). First row shows green stained coumarin-6 loaded nanoparticle; second row shows cell membrane stained red with F-actin; third row shows nuclei stained blue with Hoechst 33342 dye and fourth row shows overlay of all the three quadrants.



**Fig. S12.** Cytotoxicity of FA-DSS-dach NPs against (a) Hela cells and (b) CRL2522 (fibroblast cells) after 24, 48, and 72 h of incubation.



**Fig. S13.** (a) Viability of HeLa cells pretreated for 1 h with (a) different concentrations of *N*-ethylmaleimide (NEM) and subsequently incubated with Dulbecco's Modified Eagle Medium (DMEM) for 72 h (b) Viability of HeLa cells pretreated for 1 h with 12.5  $\mu$ M NEM and then treated with Pt(dach)Cl<sub>2</sub> or FA-DSS-Pt(dach)Cl<sub>2</sub> NPs for 72 h.



**Fig. S14.** Cytotoxicity of Pt(dach)Cl<sub>2</sub> and FA-DSS-Pt(dach)Cl<sub>2</sub> NPs against CRL2522 (fibroblast cells) after (a) 24, (b) 48, and (c) 72 h of incubation.