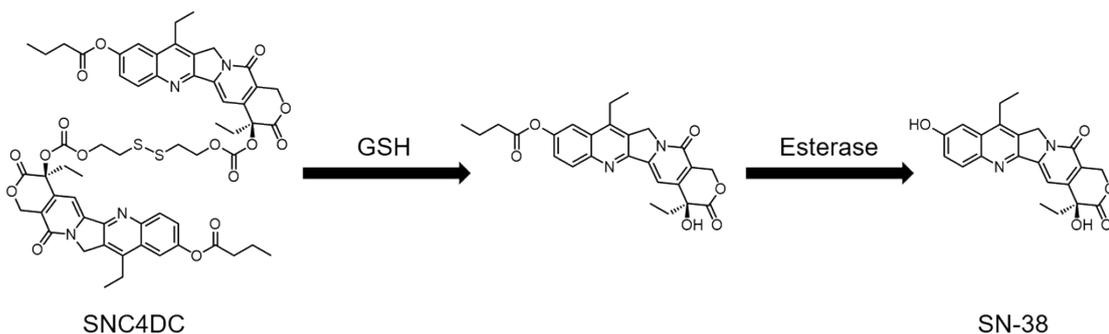


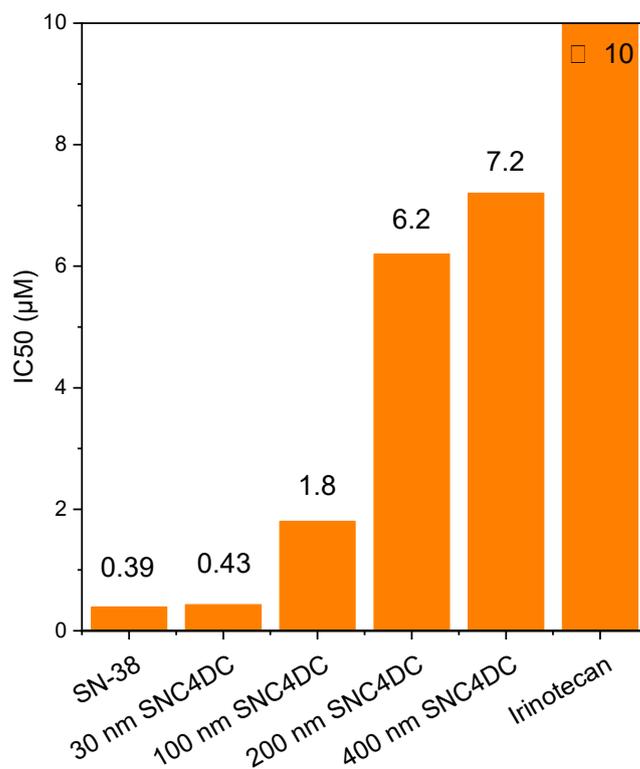
## Support Information



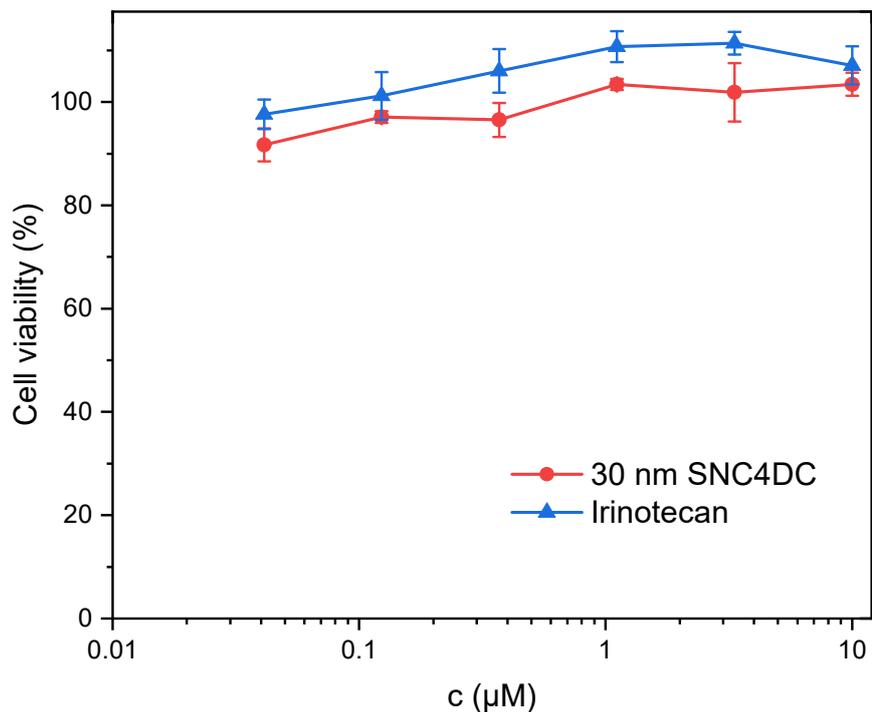
**Scheme S1.** The conversion pathway of SNC4DC into SN-38.

## *In vitro* experiment

IC<sub>50</sub> values for BxPC-3 cells, as well as the corresponding data for normal CCD-18Co cells, have been added.



**Figure S2.** BxPC-3 细胞的 IC<sub>50</sub> of SN-38 不同粒径 SNC4DC 和伊立替康

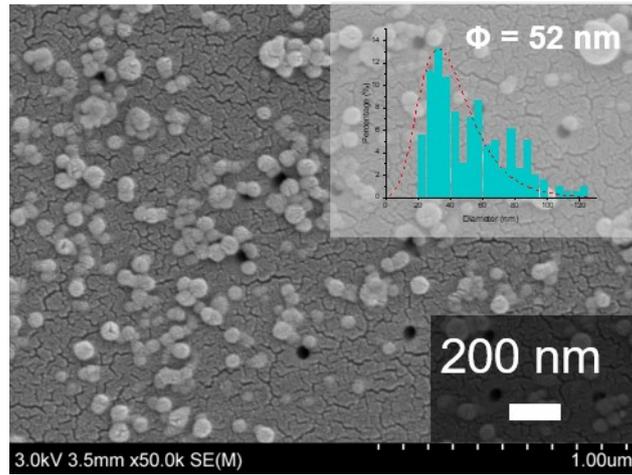


**Figure S3.** CCD-18Co cell viability curve of 30 nm SNC4DC NPDs and irinotecan.

## Experimental Section

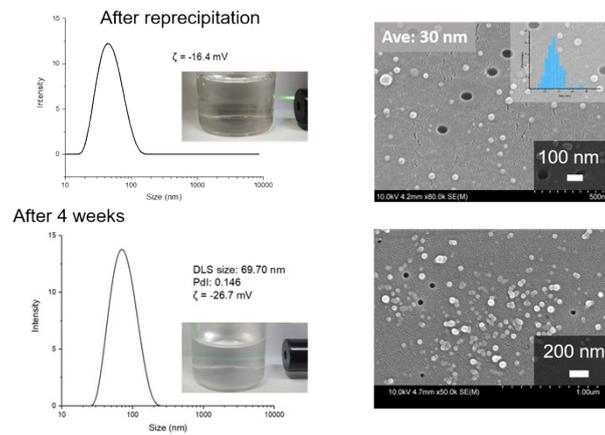
### Preparation of nanoparticle dispersions

The nanoparticle dispersions were prepared using the reprecipitation method.[9] The SNC4DC prodrug was transformed into a nanoparticle dispersion by inducing a rapid change in solubility using a 10 mM solution of SNC4DC in a good solvent (THF; 100  $\mu\text{L}$ ) and distilled water as the poor solvent. Using a 250- $\mu\text{L}$  microsyringe, the SNC4DC solution was rapidly injected into water while stirring at a speed of 1,500 rpm, followed by the cessation of stirring shortly after. The final concentration of the dispersion after reprecipitation was 0.1 mM (equal to that of SN-38).

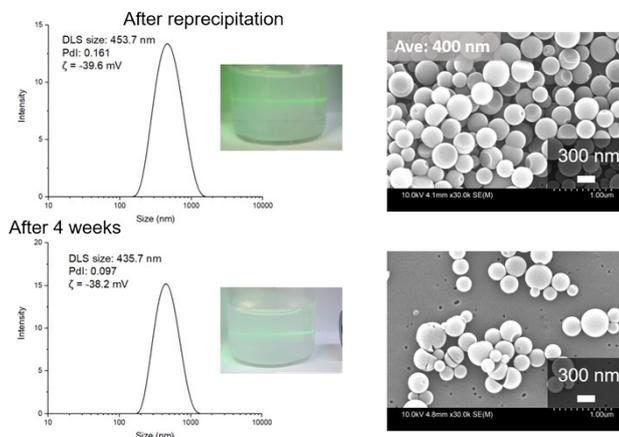


**Figure S4.** SEM image of nanoparticles with an average diameter of ~50 nm.

### Dispersion stability of nanoparticles



**Figure S5.** SEM images and DLS size distributions of 30 nm nanoparticles immediately after reprecipitation and after 4 weeks of storage.



**Figure S6.** SEM images and DLS size distributions of 400 nm nanoparticles immediately after reprecipitation and after 4 weeks of storage.

The prepared nanoparticles were evaluated using DLS (Zetasizer Nano series Nano-ZS; Malvern Instruments, Ltd.) and SEM (S-4800; Hitachi). After performing the DLS and SEM measurements, the samples were stored in a refrigerator at 4 °C for four weeks, before the measurements were repeated.