## Supporting Information

## Doxorubicin Carrier Mediated by HCOF Platform for Potent

## **Cancer Therapy**<sup>†</sup>

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## **Experimental Section**

Chemicals and Materials. 1,3,5-Tris(4-aminophenyl)benzene ( $C_{24}H_{21}N_3$ , 99%, Shanghai Macklin Biochemical Co. Ltd), 1,3,5-Benzenetricarboxaldehyde ( $C_{9}H_6O_3$ , 99.76%, Shanghai Macklin Biochemical Co. Ltd), Acetonitrile ( $C_2H_3N$ , 99.9%, Shanghai Macklin Bio chemical Co. Ltd), Folic Acid ( $C_{19}H_{19}N_7O_6$ , 97%, Shanghai Macklin Biochemical Co. Ltd), N, N-Dimethylformamide ( $C_3H_7NO$ , AR, 99.5%, Shanghai Macklin Biochemical Co. Ltd), Acetic acid ( $C_2H_4O_2$ , 99.8%, Shanghai Macklin Biochemical Co. Ltd), Ferri nitrate nonahydrate (FeH<sub>18</sub>N<sub>3</sub>O<sub>18</sub>, AR, 98.5%, Shanghai Macklin Biochemical Co. Ltd), Hydrogen peroxide ( $H_2O_2$ , 30%, Xilong Scientific).

**Characterization.** Powder X-ray diffraction (PXRD) patterns were obtained on a Riga ku MiniFlex 600 diffractometer with graphite monochromatized CuK $\alpha$  radiation ( $\lambda = 0$ . 15405 nm). The sample was scanned at a scanning rate of 5/min in the 2 $\theta$  range fro m 3 to 20° at room temperature. Transmission electron microscopy (TEM) images wer e obtained using a Talos 200 microscope (ThermoFisher Scientific, USA). A field emi ssion scanning electron microscope (Thermo Fisher, Apero 2S) was used to characteriz e the morphology of the sample. The UV-Vis adsorption spectral values were collecte on a UV-2600 spectrophotometer (Shimadzu). Fourier transform infrared spectroscopy (FT-IR) was measured on a Nicolet iS50 (ThermoFisher Scientific, USA) using the K Br tableting technique. Zeta potential data and dynamic light scattering (DLS) were ex amined employing a Zeta sizer (NanoZS, Malvern, UK). Dynamic light scattering (DL S) experiment was performed on Malvern Zeta Sizer-Nano ZS instrument at 25°C. M TT experiments were carried out using a microplate reader (Cytation5). The flow cyto metry data was obtained by BD FACSymphony A1 Cell Analyzer.

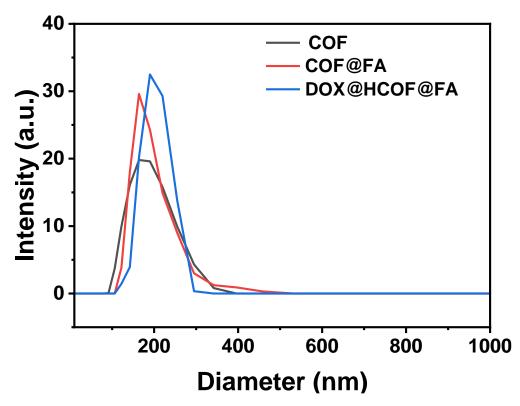


Fig. S1 The DLS measurement results for HCOF, HCOF@FA and DOX@HCOF@FA.

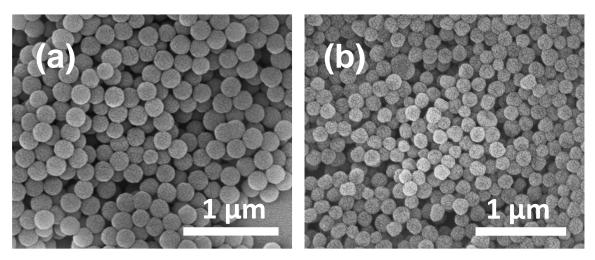
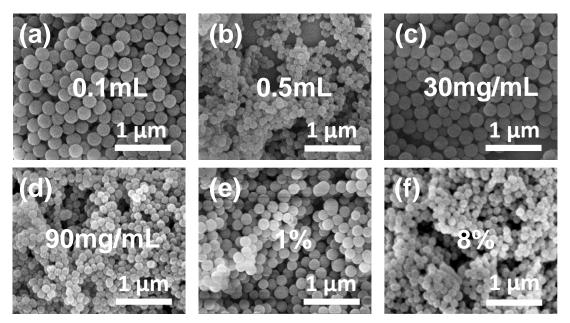


Fig. S2 SEM images of (a) COF and (b) HCOF.



**Fig. S3** SEM images of HCOF prepared with different content of (a) (b) acetic acid, (c) (d) ferric nitrate, (e) (f) hydrogen peroxide.

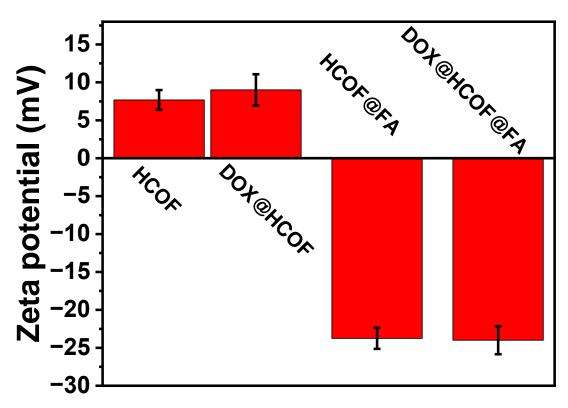


Fig. S4 Zeta potential of HCOF, HCOF@FA, DOX@HCOF and DOX@HCOF@FA.

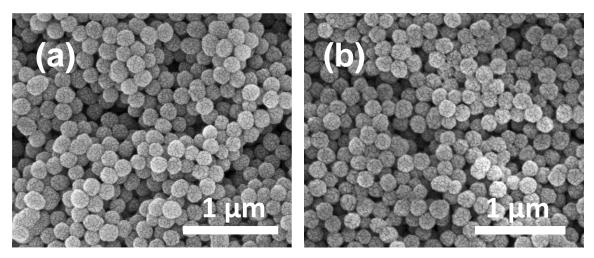


Fig. S5 SEM images of HCOF dispersed in (a) 1640 and (b) DMEM for 24 h, respectively.

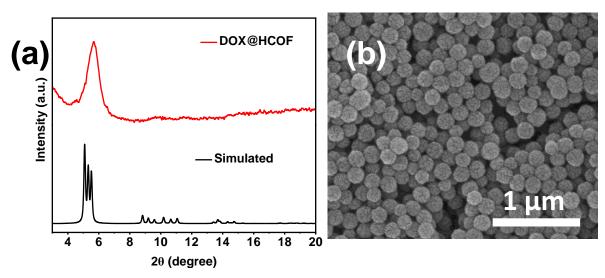


Fig. S6 (a) PXRD patten and (b) SEM image of DOX@HCOF.

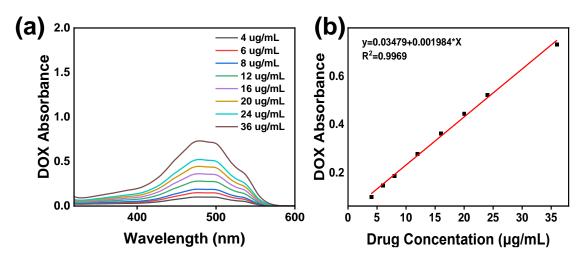


Fig. S7 (a) UV-vis absorption curve of DOX with different concentrations. (b) UV-vis absorbanceconcentration standard line of DOX.

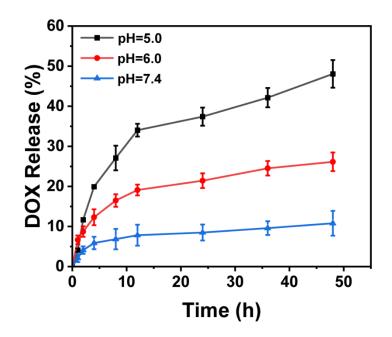
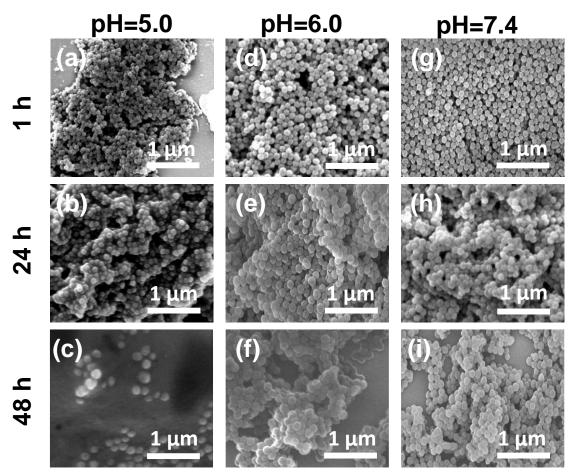


Fig. S8 Drug release profile in PBS solutions at different pH values.



**Fig. S9** SEM image of HCOF after incubating with PBS at (a-c) pH=5, (d-f) pH=6.0, (g-i) pH=7.4 for 1, 24 and 48 h, respectively.

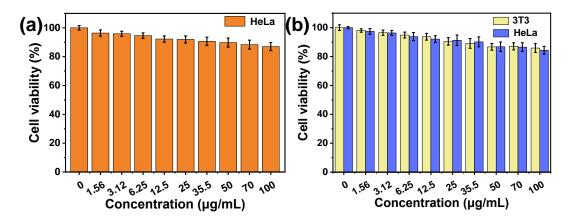


Fig. S10 (a) In vitro cell viability of HCOF@FA against HeLa cells. (b) In vitro cell viability of HCOF against 3T3 cells and HeLa cells.

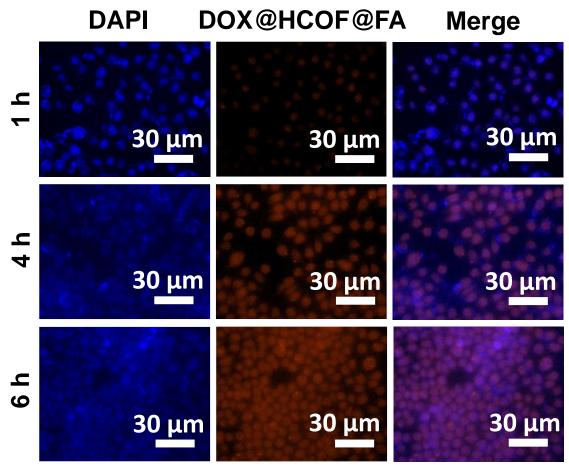


Fig. S11 CLSM images of HeLa cells incubated with DOX@HCOF@FA for different times.

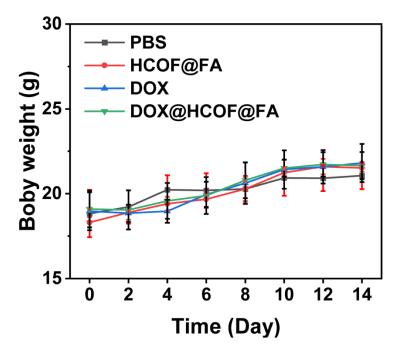


Fig. S12 Body weight changes by different treatments.