Supplementary Information

Targeting CDK4/6 in glioblastoma via in situ injection of a cellulose-based hydrogel

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Figure S1. Expression levels of CDK4 and CDK6 are higher in GBM than normal brain tissue. (A) The comparison of mRNA levels of CDK4 and CDK6 in GBM tissues (n =163) and normal brains (n = 207). (B) The immunohistochemical staining of CDK4 and CDK6 expression in GBM tissues and normal brains. The mRNA data were obtained from TCGA database and the graph in (A) were generated by GEPIA (http://gepia.cancer-pku.cn/). The protein staining of CDK4/6 of a representative example for each group were obtained from Protein Atlas (https://www.proteinatlas.org/).



Figure S2. Additional characterization of PB@PH/Cu-CNCs hydrogel. (A) The outlook and inversion test of the PB@PH/Cu-CNCs hydrogel. (B) The degradation performance of the hydrogel at pH 7. (C) The mass swelling ratio of the hydrogel at different pH 5.5 and 7.4.



Figure S3. The uncropped images of the original Western blots. (A-E) Uncroppedimages of the original WB blots related to the representative WB blots shown in theFig.4A,Fig.4D,Fig.4G,Fig.6EandFig.6J.



Figure S4. The growth inhibitory effect of PB@PH/Cu-CNCs in GSC (glioma stem cells). (A) The impact of PH/Cu-CNCs, free PB (0.1 μ M) and PB@PH/Cu-CNCs treatment on the number and phenotype of neurosphere formed by GSC23 cells. (B) Quantification of neurospheres formed in each group in (A). **, *p* < 0.01; ***, *p* < 0.001. (C) Suppression of colony formation upon treatment with PH/Cu-CNCs, free PB and PB@PH/Cu-CNCs for 9-12 days in GSC23 cells.



Figure S5. Evaluation of the in vivo toxicity of PB@PH/Cu-CNCs. H&E-stained tissue sections obtained from mice received intracranial injection of 10 μ L PH/Cu-CNCs, free PB (3.84 mg/mL) or PB@PH/Cu-CNCs (30 mg/mL) for 7 days. Scale bar = 100 μ m.