## **Supplementary Figure legends**

Figure S1. Knockout of HK2 suppresses HT29 xenograft tumor growth. A-C, average tumor volume (A), photographed xenograft tumors (B), and average tumor weight (C) of HT29 sgCtrl and HT29 sgHK2 xenografts. Data are shown as mean values  $\pm$  S.D. \*\*\*p<0.001, a significant difference between groups as indicated.

Figure S2. The structure of screened compounds.

**Figure S3.** Suppression of Akt signaling downregulates glycolysis in colorectal cancer cells. A and B, glucose consumption (A) and lactate production (B) in HCT116 and HT29 cells treated with PD98059 or wortmannin. C and D, glucose consumption (C) and lactate production (D) in HCT116 and HT29 cells transfected with siCtrl or siAkt. \*\*p<0.01, \*\*\*p<0.001, a significant difference between groups as indicated.

Fig. S1



В



С



## Fig. S2



Fig. S3

С





HCT116 HT29

D

В

