

Supplementary Methods

DeepTrio use a Siamese structure with the multiple parallel convolution module to extract various protein features from multiscale windows. Each protein was embedded as a 2D matrix and inputted into the multiscale convolution neural network which generated a group of feature maps using corresponding convolution filters. The filter outputs were activated by the rectified linear unit and yielded a set of nonlinear feature maps. The feature maps were flattened, concatenated and passed through the dense layers to get the probability vector for prediction.

Supplementary Table 1. Primers used for qPCR analyzes

| Gene | Forward Primers | Reverse primers |
|--------|--------------------------------|------------------------------|
| CDKN1A | 5' -tgtccgtcagaacccatgc-3' | 5' -aaagtcgaagttccatcgctc-3' |
| BAX | 5' -cccgagaggtcttttccgag-3' | 5' -ccagcccatgatggttctgat-3' |
| BIX | 5' -cctgcacctgct gctcaag-3' | 5' -ctccagggcagtggtcatg-3' |
| PTEN | 5' -ttgaagaccataaccaccac-3' | 5' -attaccagttcgtcccttc-3' |
| CDH1 | 5' -aaaggcccatttctctaaaacct-3' | 5' -tgcgttctctatccagaggct-3' |
| CDKN2C | 5' -ggggacctagagcaacttact-3' | 5' -cagcgcagtccttccaaat-3' |
| ADI1 | 5' -ggcctggtatatggacgacg-3' | 5' -gtcagcatccagcttccagta-3' |

Supplementary Table 2. Primers used for ChIP qRT-PCR analyzes

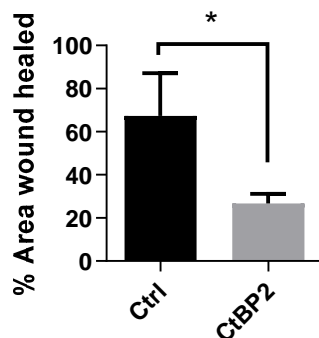
| Promoter | Forward Primers | Reverse primers |
|----------|------------------------------------|------------------------------|
| CDH1 | 5'-GTGAACCCTCAGCCAATCAG-3' | 5'-TCACAGGTGCTTTGCAGTTC-3' |
| PTEN | 5'-GGCTCAGGCGAGGGAGAT-3' | 5'-GAAGAGGCTGCACGGTTAGA-3' |
| p21 | 5'-GTGGCTCTGATTGGCTTTCTG-3' | 5'-CTGAAAACAGGCAGCCCAAG-3' |
| Bax | 5'-CCTGCCCGAAACTTCTAAAAATGG -3' | 5'-CCAATGAGCATCTCCCGATAAG-3' |

Supplementary Table 3. Protein-protein interaction prediction results for ALDOB by DeepTrio.

| Gene symbol | Description | Binding probability |
|-------------|--|---------------------|
| CTBP2 | C-terminal-binding protein 2 | 0.885 |
| CTBP1 | C-terminal-binding protein 1 | 0.326 |
| APC | Adenomatous polyposis coli protein | 0.236 |
| HIF1A | Hypoxia-inducible factor 1-alpha | 0.102 |
| HRAS | GTPase HRas | 0.085 |
| KRAS | GTPase KRas | 0.120 |
| MYB | Transcriptional activator Myb | 0.027 |
| TP53 | Cellular tumor antigen p53 | 0.120 |
| VHL | von Hippel-Lindau disease tumor suppressor | 0.071 |
| BCL2 | Apoptosis regulator Bcl-2 | 0.082 |

| | | |
|---------------|--|-------|
| BRAF | Serine/threonine-protein kinase B-raf | 0.048 |
| EGFR | Epidermal growth factor receptor | 0.188 |
| ERBB2 | Receptor tyrosine-protein kinase erbB-2 | 0.088 |
| ETV1 | ETS translocation variant 1 | 0.036 |
| ETV4 | ETS translocation variant 4 | 0.173 |
| JUN | Transcription factor AP-1 | 0.068 |
| PDGFB | Platelet-derived growth factor subunit B | 0.049 |
| RET | Proto-oncogene tyrosine-protein kinase receptor Ret | 0.242 |
| FGFR2 | Fibroblast growth factor receptor 2 | 0.019 |
| NFKB2 | Nuclear factor NF-kappa-B p100 subunit | 0.019 |
| PAX8 | Paired box protein Pax-8 | 0.020 |
| PIK3CA | Phosphatidylinositol 4,5-bisphosphate 3-kinase catalytic subunit alpha isoform | 0.054 |
| MDM2 | E3 ubiquitin-protein ligase Mdm2 | 0.092 |
| JAK2 | Tyrosine-protein kinase JAK2 | 0.045 |
| BRCA2 | Breast cancer type 2 susceptibility protein | 0.014 |
| BRCA1 | Breast cancer type 1 susceptibility protein | 0.004 |
| TSC1 | Hamartin | 0.025 |
| CDKN2C | Cyclin-dependent kinase 4 inhibitor C | 0.058 |
| WT1 | Wilms tumor protein | 0.046 |

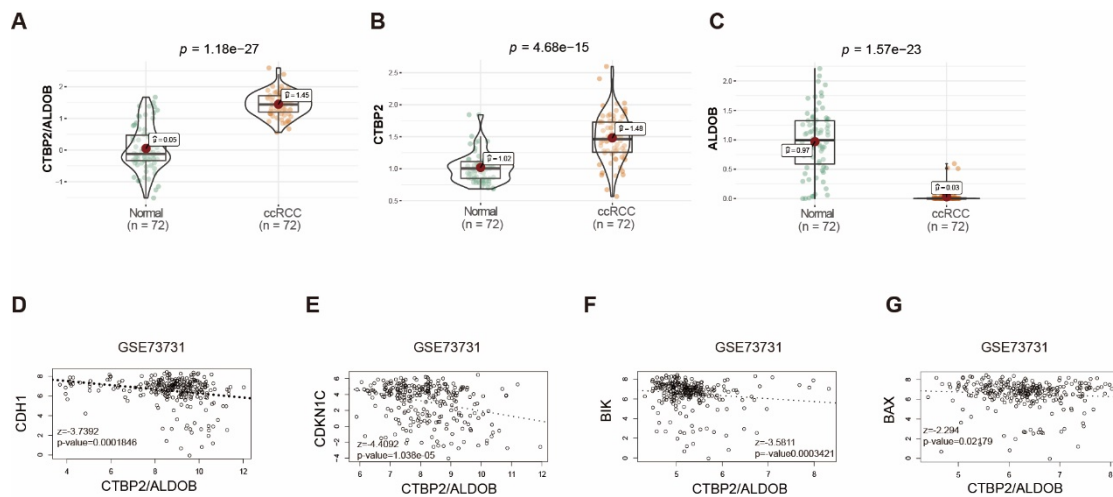
Supplementary Fig. 1 Knockdown of CtBP2 inhibits Caki-1 cell migration



Statistical analysis of the wound healing assay using Caki-1 cells transiently transfected with siCtrl or

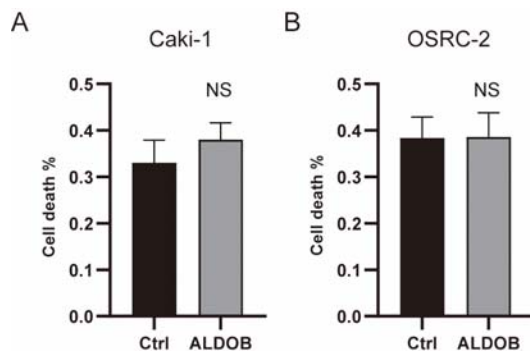
siCtBP2#1.

Supplementary Fig.2 The CtBP2-to-ALDOB expression ratio correlates with CtBP2 target gene expression and is associated with shorter survival



(A) Boxplot of the relative CTBP2-to-ALDOB expression ratio in normal kidneys and ccRCC tissues based on the GSE53757 dataset. (B-C) Boxplot of the relative CTBP2 (B) and ALDOB (C) expression in normal kidneys and ccRCC tissues based on the GSE53757 dataset. (D-G) A scatter plot was used to visualize the relationship between the CtBP2/ALDOB expression ratio and expression of target genes [CDH1 (D), CDKN2C (E), BIK (F), and BAX (G)] in the GSE73731 dataset.

Supplementary Fig. 3 Overexpression of ALDOB does not sensitize ccRCC cells to paclitaxel



(A-B) The Caki-1 (A) and OSRC-2 (B) cells were treated with 8 nM paclitaxel for 10 and 12hrs.

Percentages of dead paclitaxel-treated Caki-1 (A) and OSRC-2 (B) cells expressing Ctrl or ALDOB according to the results of the trypan blue exclusion assay. ns indicates not significant.