

Table S1 Ferroptosis-related genes

| | | | | |
|-----------|------------------|-----------------------|--------------|----------------|
| ABCC1 | ULK2 | LGMN | PIR | PEDS1 |
| ACACA | EGFR | SMPD1 | HCAR1 | ADAMTS13 |
| ACO1 | MAPK3 | MYCN | SLC16A1 | CDC25A |
| ACSF2 | MAPK1 | IFNA1 | RRM2 | CircFNDC3B |
| ACSL1 | BID | IFNA2 | NR4A1 | PPARD |
| ACSL3 | CDKN2A | IFNA4 | RPTOR | ENO3 |
| ACSL4 | SOCS1 | IFNA5 | SREBF1 | LCN2 |
| ACLS5 | CDO1 | IFNA6 | SREBF2 | MARCHF5 |
| ACSL6 | MYB | IFNA7 | FZD7 | TRIB2 |
| AIFM2 | MAPK8 | IFNA8 | P4HB | DHODH |
| AKR1C1 | MAPK9 | IFNA10 | NT5DC2 | MIR545 |
| AKR1C2 | MAPK14 | IFNA13 | BCAT2 | PDK4 |
| AKR1C3 | LINC00472 | IFNA14 | PLA2G6 | CircPVT1 |
| ALOX12 | PRKAA2 | IFNA16 | MIR424 | MIR9-3HG |
| ALOX15 | PRKAA1 | IFNA17 | PARK7 | ADIPOQ |
| ALOX5 | ELAVL1 | IFNA21 | FXN | circDTL |
| ATG5 | BAP1 | SMG9 | SUV39H1 | PTPN18 |
| ATG7 | MIR6852 | PPARG | ATF2 | ABCC5 |
| ATP5MC3 | ACVR1B | MIR335 | ACOT1 | CISD3 |
| BACH1 | TGFBR1 | SNX5 | ALDH3A2 | MS4A15 |
| CARS | EPAS1 | PAQR3 | STK11 | FURIN |
| CBS | HILPDA | MICU1 | FNDC5 | circRHBG |
| CD44 | HIF1A | TOR2A | CircIL4R | GALNT14 |
| CHAC1 | IFNG | MIR375 | CDH1 | KLHDC3 |
| CISD1 | ANO6 | MAP3K14 | NEDD4L | LINC01833 |
| CP | LPIN1 | CircKDM4C | BRD2 | circGFRA1 |
| CRYAB | HMGB1 | MIR324 | BRD3 | MAPKAP1 |
| CS | TNFAIP3 | QSOX1 | BRDT | PRR5 |
| CYBB | TLR4 | MIB2 | DECR1 | RICTOR |
| DPP4 | ATF3 | CLTRN | GLRX5 | GSTM1 |
| EMC2 | ATM | KLF2 | NCOA3 | TERT |
| FADS2 | YY1AP1 | MIR5096 | NR5A2 | circ0097009 |
| FANCD2 | EGLN2 | HOTAIR | PANX2 | TMEM161B-DT |
| FDFT1 | MIOX | H19 | RHEBP1 | circEPSTI1 |
| FTH1 | TAFAZZIN | FOXO4 | TFAP2A | MIR18A |
| FTL | MTDH | YTHDC2 | ARF6 | RARRES2 |
| FTMT | IDH1 | DDR2 | GDF15 | DUSP1 |
| G6PD | SIRT1 | SLC39A7 | ABHD12 | NCF2 |
| GCLC | FBXW7 | TRIM46 | PPP1R13L | MT3 |
| GCLM | PANX1 | KDM5A | TFAM | UBC |
| GLS2 | DNAJB6 | TRIM21 | KDM3B | ALB |
| GOT1 | LONP1 | DPEP1 | RNF113A | TXNRD1 |
| GPX4 | CD82 | CYGB | AHCY | SRXN1 |
| GSS | IL1B | IDO1 | circ-TTBK2 | GPX2 |
| HMGCR | CTSB | GSTZ1 | MIR522 | BNIP3 |
| HMOX1 | POR | GJA1 | IDH2 | OXSRI |
| HSBP1 | CYB5R1 | PGRMC1 | PPARA | SELENOS |
| HSPB1 | ELOVL5 | CIRBP | NOS2 | ANGPTL7 |
| IREB2 | FADS1 | circPSEN1 | SIAH2 | DDIT4 |
| KEAP1 | PTEN | USP11 | RELA | LOC284561 |
| LPCAT3 | NR1D1 | YAP1 | VDR | ASNS |
| MAP1LC3A | NR1D2 | MIR135B | NEDD4 | TSC22D3 |
| MAP1LC3B | TBK1 | TRIM26 | PRDX1 | DDIT3 |
| MAP1LC3C | IL6 | NDRG1 | AR | JDP2 |
| MT1G | USP7 | MIR302A | MTF1 | SLC1A4 |
| NCOA4 | miR-182-5p | ASMTL-AS1 | COPZ1 | PCK2 |
| NFE2L2 | miR-378a-3p | PIEZO1 | NUPR1 | TXNIP |
| NFS1 | ATF4 | LIFR | USP35 | VLDLR |
| NOX1 | AQP3 | PTPN6 | NEAT1 | GPT2 |
| NQO1 | AQP5 | MIR15A | PARP1 | PSAT1 |
| NRF2 | AQP8 | EGR1 | PARP2 | LURAP1L |
| OTUB1 | LINC00618 | ADAM23 | PARP3 | SLC7A5 |
| PCBP1 | MT1DP | ARHGEF26-AS1 | PARP4 | HERPUD1 |
| PCBP2 | PEX10 | CPEB1 | PARP6 | XBP1 |
| PEBP1 | AGPAT3 | COX4I2 | PARP8 | ZNF419 |
| PGD | PEX12 | lncRNA AABR07017145.1 | PARP9 | KLHL24 |
| PHKG2 | CHP1 | TIMP1 | PARP10 | TRIB3 |
| PRNP | GPAT4 | KDM6B | PARP11 | ZFP69B |
| PROM2 | BRPF1 | METTL14 | PARP12 | ATP6V1G2 |
| PTGS2 | OSBPL9 | MIB1 | PARP14 | VEGFA |
| RPL8 | INTS2 | KDM5C | PARP15 | TUBE1 |
| SAT1 | MMD | MEG3 | PARP16 | ARRDC3 |
| SAT2 | CYP4F8 | CCDC6 | PDSS2 | CEBPG |
| SLC11A2 | MLLT1 | CFL1 | TXN | SNORA16A |
| SLC1A5 | TTPA | MIR539 | SENP1 | RGS4 |
| SLC39A14 | GRIA3 | KMT2D | OIP5-AS1 | BLOC1S5-TXNDC5 |
| SLC39A8 | EPT1 | RB1 | MIR190A | LOC390705 |
| SLC3A2 | POM121L12 | HSF1 | FGF21 | EIF2S1 |
| SLC40A1 | LIG3 | SQSTM1 | CREB1 | KIM-1 |
| SLC7A11 | AEBP2 | MUC1 | CREB3 | CXCL2 |
| SQLE | AGPS | HSPA5 | CREB5 | HSD17B11 |
| STEAP3 | CDCA3 | HELLS | MIR130B | SETD1B |
| TF | PEX2 | SCD | BEX1 | MAFG |
| TFRC | PEX6 | SRC | ASAH2 | IL33 |
| TP53 | TIMM9 | STAT3 | FABP4 | HAMP |
| VDAC2 | DCAF7 | PML | AKT1S1 | DRD5 |
| VDAC3 | LCE2C | MTOR | MLST8 | DRD4 |
| ZEB1 | FAR1 | TP63 | TYRO3 | MAP3K5 |
| NOX3 | PHF21A | CDKN1A | SIRT6 | SLC2A1 |
| NOX4 | SMAD7 | MIR137 | TMSB4X | SLC2A3 |
| NOX5 | LYRM1 | ENPP2 | TMSB4Y | SLC2A6 |
| DUOX1 | AMN | FH | KIF20A | SLC2A8 |
| DUOX2 | PEX3 | CISD2 | ECH1 | SLC2A12 |
| PIK3CA | MTCH1 | MIR9-1 | circRHOT1 | GLUT13 |
| FLT3 | ACADSB | MIR9-2 | ETV4 | SLC2A14 |
| SCP2 | PVT1 | MIR9-3 | MEG8 | EIF2AK4 |
| NRAS | hsa_circ_0008367 | ISCU | VCP | TFAP2C |
| KRAS | MAP3K11 | LINC00336 | circ_0007142 | SP1 |
| HRAS | GSK3B | BRD4 | RBMS1 | HBA1 |
| TFR2 | BRD7 | PRDX6 | KDM4A | NNMT |
| SLC38A1 | SLC25A28 | MIR17 | MGST1 | PLIN4 |
| CARS1 | MFN2 | SESN2 | circKIF4A | HIC1 |
| ALOX12B | ZFAS1 | NF2 | miR-7-5p | STMN1 |
| ALOX15B | TSC1 | ARNTL | circ_0067934 | CAPG |
| ALOXE3 | TGFB1 | JUN | MPC1 | HNF4A |
| G6PDX | SNCA | CA9 | CHMP1A | NGB |
| ULK1 | SIRT3 | TMBIM4 | CAMKK2 | YWHAE |
| ATG3 | CGAS | PLIN2 | SOX2 | GABPB1 |
| ATG4D | STING1 | MIR212 | SRSF9 | AURKA |
| BECN1 | HDDC3 | Fer1HCH | PROK2 | MIR4715 |
| GABARAPL2 | MIR761 | LAMP2 | MIR4443 | RIPK1 |
| GABARAPL1 | MDM2 | ZFP36 | SIRT2 | MIR30B |
| ATG16L1 | MDM4 | CHMP5 | circRNA1615 | MMP13 |
| WIPI1 | MIR214 | CHMP6 | MIR27A | LRRFIP1 |
| WIPI2 | DLR | CAV1 | MIR670 | CBR1 |
| SNX4 | VWTR1 | GCH1 | MEF2C | |
| ATG13 | PRKCA | DAZAP1 | EZH2 | |

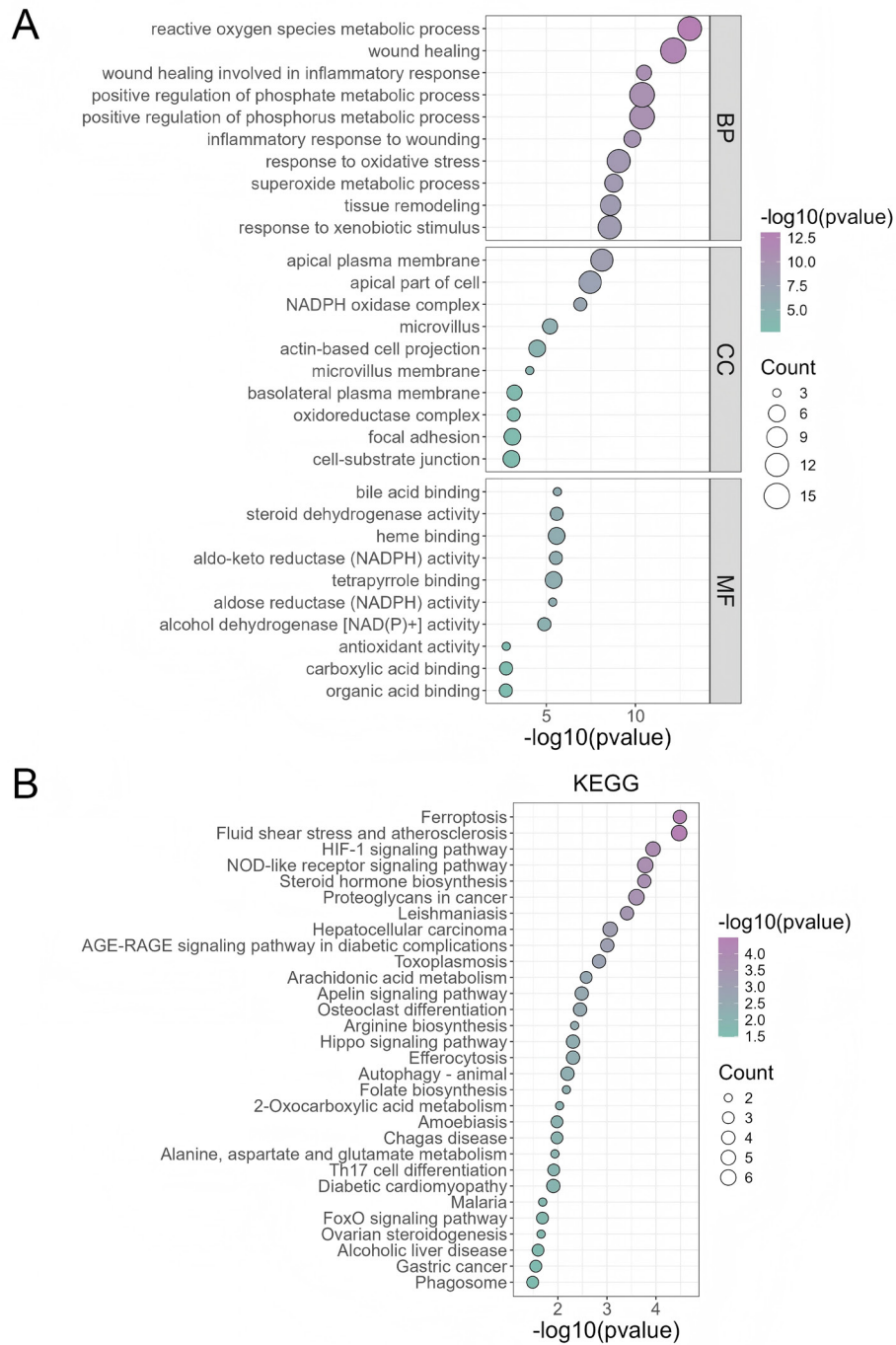


Figure S1 Functional enrichment analysis of ferroptosis-related hub genes. (A) GO enrichment analysis showing significant biological processes, cellular components, and molecular functions. (B) KEGG pathway enrichment analysis.

Table S2 Varying of transcriptional remodeling caused by virtual knockout of five model genes

| Model gene | gene | distance | Z | p.value | p.adj | logFC | |
|------------|-------------|-------------|-------------|-----------|-----------|-----------|-----------|
| AKR1C1 | GLUL | 2.35814E-05 | 1.9120157 | 5.361E-09 | 1.713E-06 | 5.0897483 | |
| | MARCKS | 2.12619E-05 | 1.8595175 | 1.428E-07 | 3.042E-05 | 4.7909923 | |
| | LMAN1 | 1.93368E-05 | 1.8119748 | 1.708E-06 | 0.0002323 | 4.5171387 | |
| | VSIG2 | 1.91434E-05 | 1.8069727 | 2.166E-06 | 0.0002323 | 4.4881399 | |
| | IGFBP2 | 1.90189E-05 | 1.803728 | 2.52E-06 | 0.0002323 | 4.4693106 | |
| | TCEAL9 | 1.90107E-05 | 1.8035142 | 2.545E-06 | 0.0002323 | 4.4680693 | |
| | TMEM98 | 1.82231E-05 | 1.7825407 | 6.496E-06 | 0.0005188 | 4.3459879 | |
| | FOXA2 | 1.69875E-05 | 1.7479738 | 2.625E-05 | 0.0018219 | 4.1433965 | |
| | CA2 | 1.69116E-05 | 1.7457799 | 2.851E-05 | 0.0018219 | 4.1304793 | |
| | AZGP1 | 1.61539E-05 | 1.7233834 | 6.401E-05 | 0.0037184 | 3.9982109 | |
| | ALDH1A1 | 1.60535E-05 | 1.7203461 | 7.107E-05 | 0.0037846 | 3.9802157 | |
| | MGST1 | 1.59212E-05 | 1.7163192 | 8.151E-05 | 0.0040064 | 3.9563371 | |
| | GPR160 | 1.52249E-05 | 1.6946302 | 0.0001648 | 0.0075224 | 3.8273081 | |
| | SH3BGRL2 | 1.45161E-05 | 1.6716387 | 0.0003279 | 0.013967 | 3.6897565 | |
| | RHOBTB3 | 1.36394E-05 | 1.6417948 | 0.0007374 | 0.0287779 | 3.5100063 | |
| | RNASE1 | 1.35976E-05 | 1.6403319 | 0.0007656 | 0.0287779 | 3.5011599 | |
| | TNFRSF12A | 1.27501E-05 | 1.6097514 | 0.0016038 | 0.0569356 | 3.3154733 | |
| | CA8 | 1.26599E-05 | 1.6063902 | 0.001731 | 0.0582172 | 3.294975 | |
| | MT1G | 1.25954E-05 | 1.6039755 | 0.0018275 | 0.05839 | 3.2802376 | |
| | MUC5AC | 1.25259E-05 | 1.6013606 | 0.0019371 | 0.0589429 | 3.2642681 | |
| | ZKSCAN1 | 1.20521E-05 | 1.5831944 | 0.0028592 | 0.0830476 | 3.1530255 | |
| | SMIM24 | 1.196E-05 | 1.5795898 | 0.0030795 | 0.0855568 | 3.1308902 | |
| | SEZ6L2 | 1.15953E-05 | 1.5650704 | 0.0041123 | 0.109489 | 3.0415175 | |
| | TRNP1 | 1.14749E-05 | 1.5601905 | 0.0045166 | 0.1112491 | 3.0114039 | |
| | KRT18 | 1.1472E-05 | 1.560075 | 0.0045266 | 0.1112491 | 3.0106903 | |
| | SMIM6 | 1.12581E-05 | 1.5512875 | 0.0053367 | 0.126302 | 2.9563644 | |
| | NEDD4L | 1.1173E-05 | 1.5477551 | 0.0056934 | 0.1299319 | 2.934491 | |
| | HSPA1A | 1.10721E-05 | 1.5435305 | 0.0061448 | 0.1353965 | 2.9083045 | |
| | TFF2 | 1.07542E-05 | 1.53 | 0.0077844 | 0.1658085 | 2.8242398 | |
| | ANXA10 | 1.04226E-05 | 1.5155091 | 0.0099022 | 0.2041134 | 2.7338758 | |
| | LIFR | 1.0353E-05 | 1.5124147 | 0.0104073 | 0.2078211 | 2.7145341 | |
| | BCAS1 | 9.86128E-06 | 1.4900289 | 0.0146743 | 0.2841478 | 2.5741395 | |
| | ANXA2 | 9.62761E-06 | 1.4790449 | 0.0171958 | 0.3231796 | 2.5049458 | |
| | KRT19 | 9.3582E-06 | 1.4660861 | 0.0205679 | 0.3755113 | 2.4230505 | |
| | SCGB2A1 | 9.18001E-06 | 1.4573341 | 0.023103 | 0.4015896 | 2.36758 | |
| | RHOB | 9.16999E-06 | 1.4568376 | 0.0232532 | 0.4015896 | 2.364429 | |
| | S100A16 | 9.12866E-06 | 1.4547844 | 0.0238821 | 0.4015961 | 2.3513952 | |
| | KRT8 | 9.05366E-06 | 1.4510378 | 0.0250609 | 0.4062482 | 2.3275927 | |
| | GALNT7 | 9.0136E-06 | 1.449025 | 0.0257111 | 0.4062482 | 2.3147953 | |
| | SELENBP1 | 8.9921E-06 | 1.4479417 | 0.026066 | 0.4062482 | 2.307905 | |
| | PGC | 8.7384E-06 | 1.4349838 | 0.0305838 | 0.4653108 | 2.2253284 | |
| | ID3 | 8.58869E-06 | 1.427181 | 0.0335535 | 0.4986209 | 2.1754644 | |
| | TMPRSS2 | 8.51204E-06 | 1.4231403 | 0.035167 | 0.49974 | 2.1496005 | |
| | ARFGEF3 | 8.51083E-06 | 1.4230763 | 0.035193 | 0.49974 | 2.1491906 | |
| | CST3 | 8.44643E-06 | 1.4196553 | 0.0366005 | 0.5084292 | 2.1272712 | |
| | ATP8B1 | 8.31656E-06 | 1.4126878 | 0.0395848 | 0.5381843 | 2.0825645 | |
| | BMP2 | 8.21486E-06 | 1.4071641 | 0.0420638 | 0.5498946 | 2.0470617 | |
| | TSC22D1 | 8.20665E-06 | 1.4067158 | 0.0422695 | 0.5498946 | 2.0441777 | |
| TM9SF3 | 8.17668E-06 | 1.4050751 | 0.0430277 | 0.5498946 | 2.0336209 | | |
| SPINK1 | 7.93067E-06 | 1.3914044 | 0.0496973 | 0.6226774 | 1.9454744 | | |
| CTSB | KRT19 | 8.40415E-06 | 2.0719064 | 1.158E-09 | 2.935E-07 | 5.2109836 | |
| | ANXA2 | 8.36556E-06 | 2.068993 | 1.378E-09 | 2.935E-07 | 5.1977036 | |
| | KRT8 | 7.94913E-06 | 2.0368027 | 8.588E-09 | 1.372E-06 | 5.050376 | |
| | KRT18 | 7.53152E-06 | 2.0030402 | 4.923E-08 | 6.292E-06 | 4.894662 | |
| | S100A6 | 7.1931E-06 | 1.9744874 | 1.899E-07 | 2.022E-05 | 4.76201 | |
| | S100A16 | 6.95199E-06 | 1.953436 | 4.794E-07 | 4.376E-05 | 4.6636345 | |
| | S100A14 | 5.66271E-06 | 1.8289722 | 4.118E-05 | 0.0032896 | 4.071766 | |
| | CLDN7 | 5.3857E-06 | 1.7991032 | 9.614E-05 | 0.0068258 | 3.9270492 | |
| | GPRC5A | 5.28479E-06 | 1.7878961 | 0.0001297 | 0.0082866 | 3.8724751 | |
| | MYOF | 5.15901E-06 | 1.7736672 | 0.000187 | 0.0108624 | 3.8029686 | |
| | ITGB4 | 4.94443E-06 | 1.7486958 | 0.0003428 | 0.0168935 | 3.6803908 | |
| | KLK10 | 4.94353E-06 | 1.7485887 | 0.0003437 | 0.0168935 | 3.6798633 | |
| | TNFRSF12A | 4.85291E-06 | 1.7377627 | 0.0004409 | 0.0201239 | 3.6264828 | |
| | KRT7 | 4.76777E-06 | 1.7274317 | 0.0005551 | 0.0236477 | 3.575408 | |
| | MYH14 | 4.41336E-06 | 1.6826623 | 0.0013935 | 0.0556517 | 3.3525352 | |
| | PRSS3 | 4.31225E-06 | 1.6693271 | 0.0017915 | 0.0673403 | 3.2856605 | |
| | CCND1 | 4.28135E-06 | 1.6651989 | 0.0019326 | 0.0686064 | 3.2649114 | |
| | S100A10 | 4.11242E-06 | 1.6421678 | 0.0029007 | 0.0975556 | 3.1487537 | |
| | CD9 | 4.03136E-06 | 1.6308295 | 0.0035074 | 0.1120603 | 3.0913169 | |
| | PLA2G16 | 3.84648E-06 | 1.60422 | 0.0053447 | 0.1581161 | 2.9558602 | |
| | ITGA2 | 3.83826E-06 | 1.6030108 | 0.0054437 | 0.1581161 | 2.9496828 | |
| | JUP | 3.81179E-06 | 1.5991049 | 0.0057737 | 0.1604096 | 2.9297144 | |
| | LGALS3 | 3.76286E-06 | 1.5918236 | 0.0064315 | 0.1712396 | 2.8924372 | |
| | SERPINB1 | 3.71435E-06 | 1.5845253 | 0.0071494 | 0.182739 | 2.8550014 | |
| | ITGA6 | 3.54946E-06 | 1.5590899 | 0.0101583 | 0.2496608 | 2.723977 | |
| | PHLDA2 | 3.52997E-06 | 1.5560182 | 0.0105797 | 0.2503866 | 2.7080947 | |
| | AHNAK2 | 3.40787E-06 | 1.5364319 | 0.0135923 | 0.3101955 | 2.6065228 | |
| | PDZK1IP1 | 3.3145E-06 | 1.5210459 | 0.016384 | 0.3610137 | 2.5263652 | |
| | CDCP1 | 3.27656E-06 | 1.5146875 | 0.0176553 | 0.3760577 | 2.4931436 | |
| | LAMC2 | 3.23944E-06 | 1.5084058 | 0.0189818 | 0.389272 | 2.4602683 | |
| | TM4SF1 | 3.22571E-06 | 1.5060669 | 0.0194941 | 0.389272 | 2.4480138 | |
| | TPM1 | 3.16194E-06 | 1.4950906 | 0.0220348 | 0.4164829 | 2.3904013 | |
| | LMO7 | 3.15576E-06 | 1.4940162 | 0.0222959 | 0.4164829 | 2.3847531 | |
| | GABRP | 3.14371E-06 | 1.4919186 | 0.0228121 | 0.4164829 | 2.3737207 | |
| | DSG2 | 3.03915E-06 | 1.473411 | 0.0277471 | 0.4925113 | 2.2761129 | |
| | SYT13 | 2.95443E-06 | 1.4580139 | 0.0323956 | 0.5594806 | 2.1945408 | |
| | C19orf33 | 2.92217E-06 | 1.4520514 | 0.0343335 | 0.5644244 | 2.1628614 | |
| | PLS1 | 2.9195E-06 | 1.4515562 | 0.0344979 | 0.5644244 | 2.160228 | |
| | MISP | 2.90615E-06 | 1.4490705 | 0.0353317 | 0.5644244 | 2.1470042 | |
| | TSPAN1 | 2.828E-06 | 1.4343197 | 0.0405657 | 0.6322318 | 2.068349 | |
| | FAT1 | 2.81323E-06 | 1.4314916 | 0.0416259 | 0.6333084 | 2.0532332 | |
| | FXYD3 | 2.78946E-06 | 1.4269152 | 0.0433807 | 0.6446572 | 2.0287478 | |
| | S100P | 2.7536E-06 | 1.4199497 | 0.0461452 | 0.6701539 | 1.9914219 | |
| | SDCBP2 | 2.7141E-06 | 1.4121839 | 0.0493618 | 0.6947744 | 1.9497239 | |
| | EZH2 | LCN2 | 1.49237E-05 | 2.5309982 | 9.861E-53 | 3.151E-50 | 7.8677932 |
| | | PDZK1IP1 | 5.56235E-06 | 1.9905813 | 1.223E-08 | 2.606E-06 | 5.0201133 |
| | | S100A10 | 3.86966E-06 | 1.8077075 | 7.403E-05 | 0.0118268 | 3.9731408 |
| | | TNFRSF12A | 3.38756E-06 | 1.7426519 | 0.000522 | 0.0667109 | 3.5892128 |
| BCAS1 | | 3.27381E-06 | 1.726123 | 0.0008002 | 0.0852228 | 3.4906648 | |
| TXN | | 3.21194E-06 | 1.7169201 | 0.0010041 | 0.0916638 | 3.4356169 | |
| STMN1 | | 3.11942E-06 | 1.7028617 | 0.0014002 | 0.1118421 | 3.3512769 | |
| HMGB2 | | 2.77934E-06 | 1.6478313 | 0.0044226 | 0.3140036 | 3.0182093 | |
| LMNA | | 2.57272E-06 | 1.6114337 | 0.0084205 | 0.5380693 | 2.7953073 | |
| TOP2A | | 2.51949E-06 | 1.6016427 | 0.0098738 | 0.5735784 | 2.7349855 | |
| ANKH | | 2.38503E-06 | 1.5760763 | 0.014585 | 0.776653 | 2.5767433 | |

Table S2 (continued)

Table S2 (continued)

| Model gene | gene | distance | Z | p.value | p.adj | logFC |
|------------|-----------|-------------|-----------|-----------|-----------|-----------|
| | HMMR | 2.305E-06 | 1.5602508 | 0.0182474 | 0.8085966 | 2.4782579 |
| | SMC4 | 2.2594E-06 | 1.5510161 | 0.0206756 | 0.8085966 | 2.420598 |
| | MKI67 | 2.2319E-06 | 1.5453686 | 0.0222721 | 0.8085966 | 2.3852667 |
| | NCAPG | 2.22167E-06 | 1.5432529 | 0.0228925 | 0.8085966 | 2.372017 |
| | CENPE | 2.18816E-06 | 1.536258 | 0.0250325 | 0.8085966 | 2.3281581 |
| | NUSAP1 | 2.1822E-06 | 1.5350053 | 0.0254303 | 0.8085966 | 2.3202949 |
| | GTSE1 | 2.174E-06 | 1.5332752 | 0.0259871 | 0.8085966 | 2.3094308 |
| | CDK1 | 2.1544E-06 | 1.5291155 | 0.0273613 | 0.8085966 | 2.2832897 |
| | KIF20B | 2.14367E-06 | 1.5268252 | 0.0281395 | 0.8085966 | 2.2688843 |
| | TUBB | 2.1206E-06 | 1.5218671 | 0.0298772 | 0.8085966 | 2.2376686 |
| | KIF11 | 2.11011E-06 | 1.5195956 | 0.0306979 | 0.8085966 | 2.2233543 |
| | BIRC5 | 2.10838E-06 | 1.519221 | 0.0308347 | 0.8085966 | 2.2209925 |
| | DSTN | 2.07143E-06 | 1.5111389 | 0.0338912 | 0.8085966 | 2.1699841 |
| | MAD2L1 | 2.06371E-06 | 1.5094333 | 0.0345619 | 0.8085966 | 2.1592056 |
| | CENPF | 2.05558E-06 | 1.5076313 | 0.0352804 | 0.8085966 | 2.1478126 |
| | CCNA2 | 2.04748E-06 | 1.5058312 | 0.0360082 | 0.8085966 | 2.1364261 |
| | ECT2 | 2.03206E-06 | 1.5023843 | 0.0374304 | 0.8085966 | 2.1146071 |
| | SFN | 2.02205E-06 | 1.5001355 | 0.0383785 | 0.8085966 | 2.1003613 |
| | ANLN | 1.98455E-06 | 1.4916215 | 0.0421146 | 0.8085966 | 2.0463484 |
| | TYMS | 1.96398E-06 | 1.486892 | 0.0442912 | 0.8085966 | 2.0162912 |
| | KIF23 | 1.96028E-06 | 1.4860353 | 0.0446933 | 0.8085966 | 2.0108426 |
| | CKS1B | 1.9521E-06 | 1.4841386 | 0.045592 | 0.8085966 | 1.9987752 |
| | DLGAP5 | 1.94531E-06 | 1.4825605 | 0.0463488 | 0.8085966 | 1.9887297 |
| | BUB1 | 1.94127E-06 | 1.4816183 | 0.0468045 | 0.8085966 | 1.9827301 |
| | PLK1 | 1.93426E-06 | 1.4799779 | 0.0476049 | 0.8085966 | 1.9722811 |
| | CEP55 | 1.93009E-06 | 1.4790013 | 0.0480856 | 0.8085966 | 1.9660587 |
| IDO1 | ANXA2 | 7.4313E-07 | 1.7804566 | 2.262E-10 | 7.226E-08 | 5.3300678 |
| | KRT19 | 7.21937E-07 | 1.7695143 | 7.204E-10 | 1.534E-07 | 5.2465856 |
| | KRT8 | 6.93661E-07 | 1.7544482 | 3.215E-09 | 5.136E-07 | 5.1313011 |
| | KRT18 | 6.7699E-07 | 1.7453 | 7.563E-09 | 9.665E-07 | 5.0611056 |
| | S100A6 | 5.99592E-07 | 1.6999319 | 3.098E-07 | 3.3E-05 | 4.7108017 |
| | S100A16 | 5.84674E-07 | 1.6905756 | 6.037E-07 | 5.511E-05 | 4.6381004 |
| | S100A14 | 4.72149E-07 | 1.6120104 | 5.586E-05 | 0.0044614 | 4.0213238 |
| | MYOF | 4.51055E-07 | 1.5953982 | 0.0001183 | 0.0083989 | 3.8894428 |
| | CLDN7 | 4.45337E-07 | 1.5907728 | 0.0001442 | 0.0092151 | 3.8526306 |
| | GPRC5A | 4.22849E-07 | 1.5720401 | 0.0003075 | 0.0178622 | 3.7031258 |
| | TNFRSF12A | 4.1258E-07 | 1.5631806 | 0.0004295 | 0.0228707 | 3.6321863 |
| | KLK10 | 4.02763E-07 | 1.5545206 | 0.0005872 | 0.0288615 | 3.5626978 |
| | ITGB4 | 3.9495E-07 | 1.54749 | 0.0007495 | 0.0330179 | 3.5061782 |
| | KRT7 | 3.93867E-07 | 1.546505 | 0.0007751 | 0.0330179 | 3.4982521 |
| | PRSS3 | 3.77475E-07 | 1.5312923 | 0.0012746 | 0.0509024 | 3.3755989 |
| | CCND1 | 3.53901E-07 | 1.5083196 | 0.002524 | 0.0948728 | 3.1895261 |
| | MYH14 | 3.49838E-07 | 1.5042203 | 0.0028286 | 0.1004146 | 3.1562136 |
| | CD9 | 3.40216E-07 | 1.4943339 | 0.0036884 | 0.124046 | 3.075737 |
| | PLA2G16 | 3.28828E-07 | 1.4822987 | 0.0050088 | 0.1600324 | 2.9775079 |
| | JUP | 3.10466E-07 | 1.462065 | 0.0080551 | 0.2327986 | 2.8111712 |
| | AHNAK2 | 3.09844E-07 | 1.4613606 | 0.0081826 | 0.2327986 | 2.805925 |
| | S100A10 | 3.08901E-07 | 1.4602902 | 0.0083793 | 0.2327986 | 2.7971296 |
| | PHLDA2 | 3.07184E-07 | 1.4583328 | 0.0087484 | 0.2329269 | 2.7810401 |
| | SERPINB1 | 3.04317E-07 | 1.4550434 | 0.009397 | 0.240187 | 2.7539845 |
| | PDZK1IP1 | 3.01201E-07 | 1.4514365 | 0.0101501 | 0.2494586 | 2.724292 |
| | ITGA2 | 2.99613E-07 | 1.4495849 | 0.0105543 | 0.2497851 | 2.7090389 |
| | ITGA6 | 2.93826E-07 | 1.4427611 | 0.0121509 | 0.2773002 | 2.6527667 |
| | LGALS3 | 2.91002E-07 | 1.4393858 | 0.0130052 | 0.2865625 | 2.6248974 |
| | TM4SF1 | 2.69954E-07 | 1.4132443 | 0.0212235 | 0.4520599 | 2.408263 |
| | LAMC2 | 2.58835E-07 | 1.3986734 | 0.0271684 | 0.5580937 | 2.2869039 |
| | GABRP | 2.56955E-07 | 1.3961527 | 0.0283045 | 0.5580937 | 2.2658643 |
| | CDCP1 | 2.5612E-07 | 1.3950284 | 0.0288217 | 0.5580937 | 2.2564761 |
| | TPM1 | 2.47258E-07 | 1.3828803 | 0.0348344 | 0.6421082 | 2.1548667 |
| | LMO7 | 2.46802E-07 | 1.3822455 | 0.0351702 | 0.6421082 | 2.149549 |
| | TIMP1 | 2.35466E-07 | 1.3660845 | 0.0444693 | 0.7893309 | 2.0138736 |
| | GLUL | 2.32109E-07 | 1.3611627 | 0.0475922 | 0.7932063 | 1.9724452 |
| | FAT1 | 2.32016E-07 | 1.3610254 | 0.0476814 | 0.7932063 | 1.9712883 |
| | SYT13 | 2.31259E-07 | 1.3599061 | 0.0484117 | 0.7932063 | 1.9618595 |
| TIMP1 | KRT19 | 2.16965E-05 | 2.0183624 | 2.394E-10 | 7.65E-08 | 5.3260687 |
| | ANXA2 | 2.1267E-05 | 2.0071755 | 5.357E-10 | 1.141E-07 | 5.2683819 |
| | KRT8 | 2.02775E-05 | 1.9806332 | 3.231E-09 | 5.161E-07 | 5.1309091 |
| | KRT18 | 1.9462E-05 | 1.9578985 | 1.336E-08 | 1.708E-06 | 5.0124759 |
| | S100A6 | 1.81275E-05 | 1.9188397 | 1.211E-07 | 1.29E-05 | 4.8075157 |
| | S100A16 | 1.76022E-05 | 1.9027739 | 2.77E-07 | 2.529E-05 | 4.7226576 |
| | S100A14 | 1.43239E-05 | 1.791899 | 2.897E-05 | 0.0023137 | 4.1280024 |
| | CLDN7 | 1.32954E-05 | 1.752537 | 0.0001039 | 0.0073803 | 3.9129966 |
| | TNFRSF12A | 1.28755E-05 | 1.7357053 | 0.0001708 | 0.0107775 | 3.8204175 |
| | MYOF | 1.27851E-05 | 1.7320183 | 0.0001898 | 0.0107775 | 3.8000857 |
| | GPRC5A | 1.27295E-05 | 1.729739 | 0.0002024 | 0.0107775 | 3.7875076 |
| | KLK10 | 1.24741E-05 | 1.7191595 | 0.0002711 | 0.0133256 | 3.7290319 |
| | ITGB4 | 1.21025E-05 | 1.7034213 | 0.0004109 | 0.0187561 | 3.6417565 |
| | KRT7 | 1.18139E-05 | 1.6909093 | 0.0005632 | 0.0239941 | 3.5721257 |
| | PRSS3 | 1.08343E-05 | 1.6463526 | 0.0015628 | 0.062414 | 3.3223716 |
| | MYH14 | 1.06534E-05 | 1.6377429 | 0.0018712 | 0.070334 | 3.2737855 |
| | CCND1 | 1.0423E-05 | 1.6265913 | 0.0023447 | 0.0832377 | 3.2106962 |
| | CD9 | 9.72313E-06 | 1.5913504 | 0.0045343 | 0.1524961 | 3.0101357 |
| | PLA2G16 | 9.46049E-06 | 1.5775543 | 0.00575 | 0.1797386 | 2.9311237 |
| | ITGA2 | 9.43033E-06 | 1.5759487 | 0.0059069 | 0.1797386 | 2.9219097 |
| | S100A10 | 9.33736E-06 | 1.5709713 | 0.0064153 | 0.1863343 | 2.8933224 |
| | PDZK1IP1 | 9.12957E-06 | 1.5596889 | 0.0076964 | 0.213826 | 2.8283862 |
| | JUP | 8.99698E-06 | 1.5523725 | 0.0086293 | 0.2228388 | 2.7861744 |
| | LGALS3 | 8.98501E-06 | 1.5517078 | 0.0087183 | 0.2228388 | 2.7823352 |
| | SERPINB1 | 8.89401E-06 | 1.5466259 | 0.0094222 | 0.2297915 | 2.7529628 |
| | PHLDA2 | 8.8586E-06 | 1.5446362 | 0.0097095 | 0.2297915 | 2.7414522 |
| | AHNAK2 | 8.53291E-06 | 1.5260038 | 0.0127409 | 0.2907652 | 2.6333702 |
| | ITGA6 | 8.4755E-06 | 1.5226555 | 0.0133545 | 0.2942603 | 2.6138917 |
| | LAMC2 | 7.728E-06 | 1.4771552 | 0.0240731 | 0.5127564 | 2.3474844 |
| | CDCP1 | 7.66751E-06 | 1.4733075 | 0.0252014 | 0.5194743 | 2.3248085 |
| | TM4SF1 | 7.60895E-06 | 1.4695577 | 0.026337 | 0.5259162 | 2.3026874 |
| | GABRP | 7.56421E-06 | 1.4666764 | 0.0272339 | 0.5273479 | 2.2856743 |
| | LMO7 | 7.409E-06 | 1.4565626 | 0.0305535 | 0.5742265 | 2.2258533 |
| | TPM1 | 7.33627E-06 | 1.4517595 | 0.0322251 | 0.5883383 | 2.1973878 |
| | SYT13 | 7.07466E-06 | 1.434134 | 0.0389006 | 0.6904857 | 2.0926144 |
| | DSG2 | 6.9928E-06 | 1.4285024 | 0.041217 | 0.7118291 | 2.0590333 |
| | C19orf33 | 6.94699E-06 | 1.4253259 | 0.0425635 | 0.7157396 | 2.0400693 |
| | MISP | 6.90914E-06 | 1.4226875 | 0.043704 | 0.7160733 | 2.0243058 |
| | TSPAN1 | 6.85234E-06 | 1.4187045 | 0.0454638 | 0.7262846 | 2.000487 |
| | PLS1 | 6.74401E-06 | 1.4110281 | 0.0489858 | 0.7634612 | 1.9545092 |

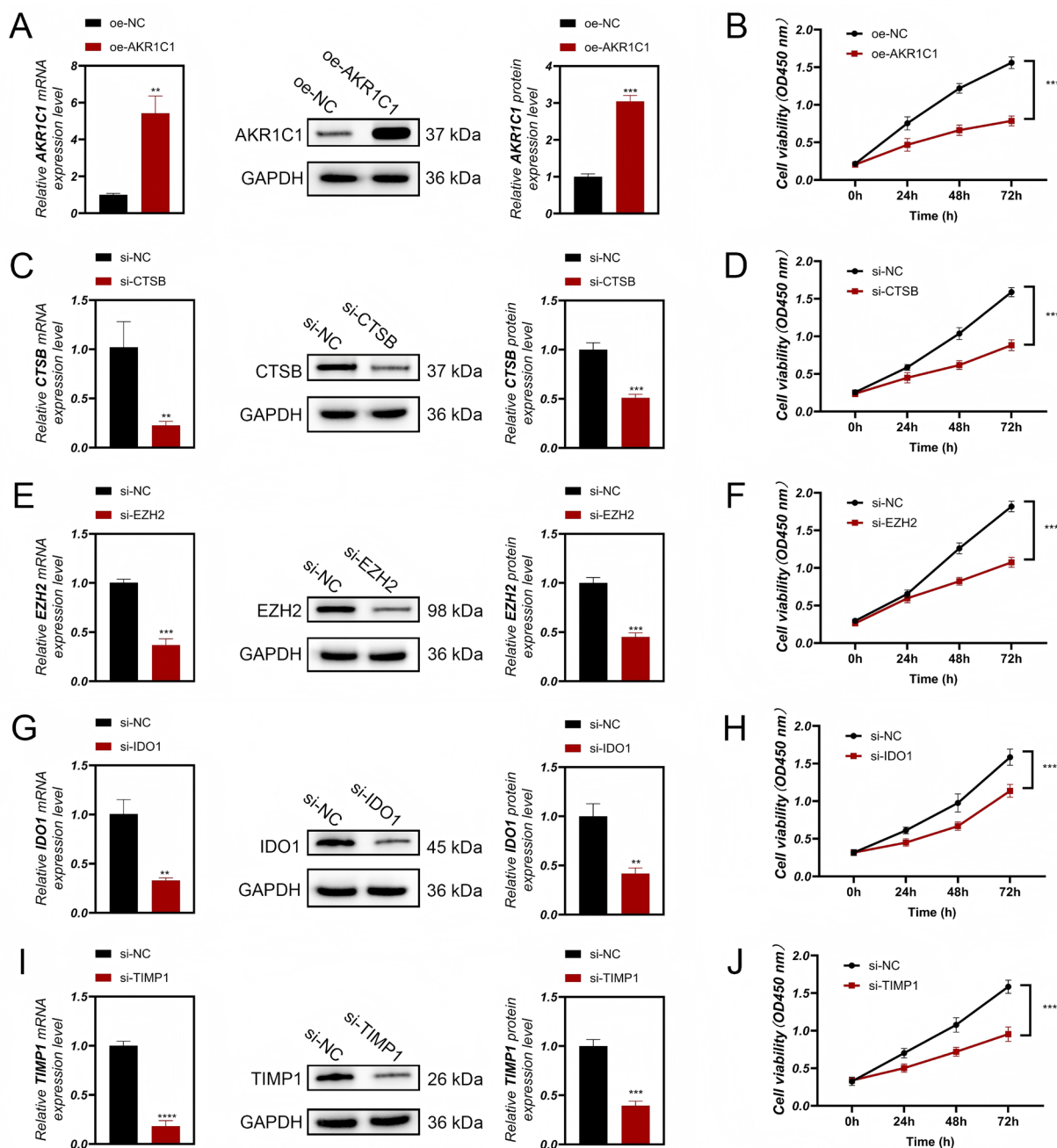


Figure S2 Validation of gene overexpression or knockdown in HGC-27 cells and effects on proliferation. (A) qRT-PCR and Western blot confirmed effective overexpression of AKR1C1 (oe-AKR1C1) and knockdown of CTSB, EZH2, IDO1, and TIMP1 (siRNAs). (B) CCK-8 assays showed that AKR1C1 overexpression or knockdown of the other four genes significantly inhibited cell proliferation. **, $P < 0.01$; ***, $P < 0.001$.