

Single-gene prognostic signatures for advanced stage serous ovarian cancer based on 1257 patient samples

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The process of principal component regression

Firstly, we conducted principal component analysis for 341 genes in EB dataset.

Secondly, the scores of principal components in each sample were calculated by multiplying loadings and gene expression values.

Thirdly, we conducted cox regression for different number of principal components.

The third principal component was not statistically significant. So, we selected first 2 components.

R code for principal component regression in this study

```
data=read.csv("G:\\dataForPCR.csv")
data2=data[,4:ncol(data)]
#principle component analysis
pca<-prcomp(data2)
#The loadings of 341 genes in the first 2 components
loadings=pca$rotation[,1:2]
#cox regression for first 2 components
pc12=data.frame(pca$x[,1:2],status=data[,2],time=data[,3])
library(survival)
coxph(Surv(time, status)~PC1+PC2,data=pc12)
```

Supplementary Table S1. The detailed information of genes associated with OS in this study.

| No. | Gene | P | FDR | C | HR(CI) |
|--------------------------------|---------|----------|----------|-------|-----------------|
| Positively OS-associated genes | | | | | |
| 1 | GMPR | 1.27E-07 | 5.30E-04 | -0.18 | 0.84(0.78-0.89) |
| 2 | ALDH5A1 | 5.71E-07 | 1.23E-03 | -0.22 | 0.8(0.74-0.88) |
| 3 | PCK2 | 2.75E-06 | 3.27E-03 | -0.29 | 0.75(0.66-0.84) |
| 4 | GJB1 | 4.04E-06 | 3.27E-03 | -0.26 | 0.77(0.69-0.86) |
| 5 | DHRS4 | 4.38E-06 | 3.27E-03 | -0.33 | 0.72(0.62-0.83) |
| 6 | SNRPA1 | 9.36E-06 | 4.24E-03 | -0.25 | 0.78(0.7-0.87) |
| 7 | PRIM2 | 1.06E-05 | 4.24E-03 | -0.29 | 0.75(0.65-0.85) |
| 8 | EMC9 | 1.80E-05 | 4.47E-03 | -0.31 | 0.73(0.64-0.85) |
| 9 | STOML2 | 2.14E-05 | 4.47E-03 | -0.25 | 0.78(0.69-0.87) |
| 10 | CAAP1 | 2.16E-05 | 6.38E-03 | -0.23 | 0.79(0.71-0.88) |
| 11 | MCM5 | 2.57E-05 | 8.43E-03 | -0.20 | 0.82(0.75-0.9) |
| 12 | PSME2P2 | 2.69E-05 | 8.43E-03 | -0.26 | 0.77(0.69-0.87) |
| 13 | IL6R | 2.87E-05 | 8.43E-03 | -0.16 | 0.85(0.79-0.92) |
| 14 | PLAA | 3.52E-05 | 1.78E-02 | -0.29 | 0.75(0.66-0.86) |
| 15 | CD38 | 3.76E-05 | 1.78E-02 | -0.21 | 0.81(0.73-0.89) |
| 16 | SLC37A4 | 4.54E-05 | 1.78E-02 | -0.29 | 0.75(0.65-0.86) |
| 17 | CXCL11 | 5.08E-05 | 1.78E-02 | -0.10 | 0.9(0.86-0.95) |
| 18 | POLD3 | 5.49E-05 | 1.78E-02 | -0.29 | 0.75(0.65-0.86) |
| 19 | FOXJ1 | 5.94E-05 | 1.78E-02 | -0.18 | 0.83(0.76-0.91) |
| 20 | OXLD1 | 6.78E-05 | 1.78E-02 | -0.25 | 0.78(0.68-0.88) |
| 21 | TRIM27 | 8.83E-05 | 2.15E-02 | -0.25 | 0.78(0.69-0.88) |
| 22 | CCL13 | 9.18E-05 | 2.15E-02 | -0.32 | 0.72(0.62-0.85) |
| 23 | ZNF165 | 1.06E-04 | 2.15E-02 | -0.20 | 0.82(0.74-0.91) |
| 24 | CYB561 | 1.11E-04 | 2.15E-02 | -0.26 | 0.77(0.68-0.88) |
| 25 | CYBA | 1.15E-04 | 2.15E-02 | -0.19 | 0.83(0.76-0.91) |
| 26 | CNOT4 | 1.16E-04 | 2.15E-02 | -0.28 | 0.75(0.65-0.87) |
| 27 | TESK1 | 1.22E-04 | 2.15E-02 | -0.31 | 0.73(0.63-0.86) |
| 28 | ACOT13 | 1.25E-04 | 2.15E-02 | -0.22 | 0.8(0.72-0.9) |
| 29 | SDF2L1 | 1.35E-04 | 2.15E-02 | -0.20 | 0.82(0.74-0.91) |
| 30 | WDR77 | 1.36E-04 | 2.15E-02 | -0.26 | 0.77(0.67-0.88) |
| 31 | SLC27A2 | 1.39E-04 | 2.15E-02 | -0.30 | 0.74(0.63-0.86) |
| 32 | GSTZ1 | 1.43E-04 | 2.15E-02 | -0.22 | 0.8(0.71-0.9) |
| 33 | FCHSD2 | 1.50E-04 | 2.15E-02 | -0.25 | 0.78(0.68-0.88) |
| 34 | ST6GAL1 | 1.55E-04 | 2.15E-02 | -0.15 | 0.86(0.79-0.93) |
| 35 | VCP | 1.57E-04 | 2.36E-02 | -0.22 | 0.8(0.72-0.9) |
| 36 | RMI1 | 1.86E-04 | 2.36E-02 | -0.21 | 0.81(0.73-0.91) |
| 37 | FEN1 | 1.86E-04 | 2.36E-02 | -0.20 | 0.82(0.74-0.91) |
| 38 | PRPS2 | 1.87E-04 | 2.36E-02 | -0.18 | 0.84(0.76-0.92) |
| 39 | NFX1 | 2.03E-04 | 2.36E-02 | -0.25 | 0.78(0.68-0.89) |
| 40 | GALNT6 | 2.13E-04 | 2.36E-02 | -0.15 | 0.86(0.8-0.93) |

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|----|-----------|----------|----------|-------|-----------------|
| 41 | RCL1 | 2.14E-04 | 2.36E-02 | -0.29 | 0.74(0.64-0.87) |
| 42 | GCNT1 | 2.15E-04 | 2.36E-02 | -0.26 | 0.77(0.67-0.89) |
| 43 | FN3KRP | 2.16E-04 | 2.36E-02 | -0.28 | 0.76(0.66-0.88) |
| 44 | SLC5A1 | 2.36E-04 | 2.36E-02 | -0.16 | 0.86(0.79-0.93) |
| 45 | SPDEF | 2.46E-04 | 2.36E-02 | -0.16 | 0.85(0.78-0.93) |
| 46 | TRIM26 | 2.50E-04 | 2.36E-02 | -0.29 | 0.75(0.64-0.88) |
| 47 | CHAF1B | 2.55E-04 | 2.36E-02 | -0.28 | 0.76(0.65-0.88) |
| 48 | CASP8AP2 | 2.57E-04 | 2.36E-02 | -0.22 | 0.81(0.72-0.9) |
| 49 | PIR | 2.58E-04 | 2.36E-02 | -0.14 | 0.87(0.81-0.94) |
| 50 | GALNT12 | 2.64E-04 | 3.17E-02 | -0.16 | 0.86(0.79-0.93) |
| 51 | POLA2 | 2.88E-04 | 3.17E-02 | -0.29 | 0.75(0.64-0.88) |
| 52 | GZMB | 2.91E-04 | 3.17E-02 | -0.15 | 0.86(0.79-0.93) |
| 53 | MCM3 | 3.03E-04 | 3.17E-02 | -0.19 | 0.82(0.74-0.92) |
| 54 | STXBP6 | 3.08E-04 | 3.17E-02 | -0.10 | 0.91(0.86-0.96) |
| 55 | MCAT | 3.13E-04 | 3.17E-02 | -0.28 | 0.76(0.65-0.88) |
| 56 | C9orf116 | 3.20E-04 | 3.17E-02 | -0.22 | 0.8(0.71-0.91) |
| 57 | SYK | 3.21E-04 | 3.17E-02 | -0.17 | 0.85(0.77-0.93) |
| 58 | BCAP31 | 3.35E-04 | 3.17E-02 | -0.22 | 0.8(0.71-0.9) |
| 59 | COX7A2P2 | 3.40E-04 | 3.17E-02 | -0.24 | 0.78(0.69-0.9) |
| 60 | BRCC3 | 3.40E-04 | 3.17E-02 | -0.22 | 0.81(0.72-0.91) |
| 61 | SDR39U1 | 3.45E-04 | 3.17E-02 | -0.26 | 0.77(0.67-0.89) |
| 62 | ZNHIT2 | 3.56E-04 | 3.17E-02 | -0.31 | 0.73(0.62-0.87) |
| 63 | LRRK1 | 3.68E-04 | 3.17E-02 | -0.27 | 0.76(0.66-0.89) |
| 64 | WDR3 | 3.89E-04 | 3.55E-02 | -0.21 | 0.81(0.72-0.91) |
| 65 | SIRT5 | 4.00E-04 | 3.55E-02 | -0.20 | 0.82(0.73-0.91) |
| 66 | COA4 | 4.00E-04 | 3.55E-02 | -0.22 | 0.8(0.71-0.91) |
| 67 | EFTUD1 | 4.12E-04 | 3.55E-02 | -0.26 | 0.77(0.66-0.89) |
| 68 | FAM86B2 | 4.26E-04 | 3.55E-02 | -0.40 | 0.67(0.53-0.84) |
| 69 | WDR45B | 4.44E-04 | 3.55E-02 | -0.21 | 0.81(0.72-0.91) |
| 70 | CIB1 | 4.47E-04 | 3.55E-02 | -0.20 | 0.82(0.74-0.92) |
| 71 | NT5C | 4.55E-04 | 4.05E-02 | -0.34 | 0.71(0.59-0.86) |
| 72 | ENDOG | 4.60E-04 | 4.05E-02 | -0.20 | 0.82(0.73-0.91) |
| 73 | UBD | 4.96E-04 | 4.05E-02 | -0.07 | 0.93(0.89-0.97) |
| 74 | POLG | 4.98E-04 | 4.05E-02 | -0.28 | 0.76(0.65-0.89) |
| 75 | FOCAD | 5.05E-04 | 4.05E-02 | -0.21 | 0.81(0.73-0.91) |
| 76 | AADAC | 5.22E-04 | 4.05E-02 | -0.13 | 0.88(0.81-0.94) |
| 77 | SLC25A1 | 5.42E-04 | 4.05E-02 | -0.20 | 0.82(0.73-0.92) |
| 78 | WARS | 5.51E-04 | 4.05E-02 | -0.16 | 0.85(0.78-0.93) |
| 79 | SYNGR2 | 5.59E-04 | 4.05E-02 | -0.18 | 0.83(0.75-0.92) |
| 80 | CREB3 | 5.79E-04 | 4.05E-02 | -0.26 | 0.77(0.67-0.9) |
| 81 | GPRC5C | 6.26E-04 | 4.05E-02 | -0.15 | 0.86(0.79-0.94) |
| 82 | DNAJA1 | 6.28E-04 | 4.05E-02 | -0.19 | 0.82(0.74-0.92) |
| 83 | LINC00338 | 6.40E-04 | 4.05E-02 | -0.23 | 0.8(0.7-0.91) |
| 84 | MAK | 6.67E-04 | 4.05E-02 | -0.19 | 0.83(0.74-0.92) |

| Positively OS-associated genes | | | | | |
|--------------------------------|----------------|----------|----------|-------|-----------------|
| 85 | TBCC | 7.10E-04 | 4.05E-02 | -0.25 | 0.78(0.68-0.9) |
| 86 | TAP2 | 7.29E-04 | 4.05E-02 | -0.15 | 0.86(0.79-0.94) |
| 87 | KRT18P38 | 7.59E-04 | 4.05E-02 | -0.53 | 0.59(0.43-0.8) |
| 88 | EFHC2 | 7.69E-04 | 4.05E-02 | -0.14 | 0.87(0.8-0.94) |
| 89 | GMPR2 | 7.70E-04 | 4.05E-02 | -0.21 | 0.81(0.72-0.92) |
| 90 | LRIF1 | 7.75E-04 | 4.37E-02 | -0.21 | 0.81(0.72-0.92) |
| 91 | TIPIN | 7.84E-04 | 4.37E-02 | -0.22 | 0.8(0.71-0.91) |
| 92 | MRPS11 | 7.89E-04 | 4.37E-02 | -0.24 | 0.79(0.69-0.91) |
| 93 | SLC7A11 | 8.18E-04 | 4.37E-02 | -0.12 | 0.89(0.83-0.95) |
| 94 | SENP6 | 8.26E-04 | 4.37E-02 | -0.20 | 0.82(0.73-0.92) |
| 95 | GCH1 | 8.33E-04 | 4.37E-02 | -0.14 | 0.87(0.81-0.95) |
| 96 | SURF2 | 8.37E-04 | 4.37E-02 | -0.30 | 0.74(0.62-0.88) |
| 97 | C6orf165 | 8.50E-04 | 4.37E-02 | -0.20 | 0.82(0.72-0.92) |
| 98 | FAH | 8.93E-04 | 4.37E-02 | -0.14 | 0.87(0.8-0.94) |
| 99 | TDP2 | 8.99E-04 | 4.37E-02 | -0.18 | 0.83(0.75-0.93) |
| 100 | MAGEF1 | 9.05E-04 | 4.37E-02 | -0.21 | 0.81(0.72-0.92) |
| 101 | ZNF440 | 9.06E-04 | 4.37E-02 | -0.26 | 0.77(0.66-0.9) |
| 102 | KIAA0020 | 9.19E-04 | 4.37E-02 | -0.21 | 0.81(0.72-0.92) |
| 103 | TAP1 | 9.19E-04 | 4.37E-02 | -0.11 | 0.9(0.84-0.96) |
| 104 | SDHAP3 | 9.20E-04 | 4.37E-02 | -0.24 | 0.79(0.69-0.91) |
| 105 | FRYL | 9.23E-04 | 4.37E-02 | -0.23 | 0.79(0.69-0.91) |
| 106 | LRIG1 | 9.29E-04 | 4.37E-02 | -0.11 | 0.89(0.84-0.96) |
| 107 | THOP1 | 9.37E-04 | 4.37E-02 | -0.34 | 0.71(0.58-0.87) |
| 108 | HLA-DOB | 9.52E-04 | 4.37E-02 | -0.19 | 0.83(0.74-0.93) |
| 109 | NAT10 | 9.60E-04 | 4.37E-02 | -0.23 | 0.79(0.69-0.91) |
| 110 | LPAR2 | 9.60E-04 | 4.37E-02 | -0.24 | 0.79(0.68-0.91) |
| Negatively OS-associated genes | | | | | |
| 1 | HSPB7 | 1.08E-07 | 5.30E-04 | 0.54 | 1.71(1.41-2.09) |
| 2 | ASAP3 | 1.68E-07 | 8.69E-04 | 0.35 | 1.41(1.24-1.61) |
| 3 | AXL | 2.60E-07 | 1.23E-03 | 0.26 | 1.3(1.17-1.43) |
| 4 | CILP | 4.44E-07 | 1.23E-03 | 0.15 | 1.16(1.1-1.23) |
| 5 | MFAP4 | 6.52E-07 | 3.27E-03 | 0.17 | 1.19(1.11-1.27) |
| 6 | LRP1 | 1.07E-06 | 3.27E-03 | 0.35 | 1.42(1.23-1.63) |
| 7 | TMEM256-PLSCR3 | 1.60E-06 | 3.27E-03 | 0.35 | 1.42(1.23-1.63) |
| 8 | LUZP1 | 1.81E-06 | 3.27E-03 | 0.41 | 1.5(1.27-1.77) |
| 9 | RAB11FIP5 | 2.93E-06 | 3.27E-03 | 0.40 | 1.5(1.26-1.77) |
| 10 | ZFHX4 | 3.22E-06 | 3.27E-03 | 0.27 | 1.31(1.17-1.48) |
| 11 | EHD2 | 3.61E-06 | 3.27E-03 | 0.31 | 1.37(1.2-1.56) |
| 12 | HP1BP3 | 4.36E-06 | 3.27E-03 | 0.27 | 1.31(1.17-1.47) |
| 13 | LRRN3 | 4.46E-06 | 4.24E-03 | 0.33 | 1.39(1.21-1.6) |
| 14 | TMEM204 | 6.38E-06 | 4.24E-03 | 0.31 | 1.36(1.19-1.55) |
| 15 | MAL | 6.41E-06 | 4.24E-03 | 0.09 | 1.09(1.05-1.13) |
| 16 | CRYAB | 6.97E-06 | 4.24E-03 | 0.12 | 1.13(1.07-1.19) |
| 17 | SVIL | 7.49E-06 | 4.24E-03 | 0.20 | 1.22(1.12-1.33) |

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|----|-------------|----------|----------|------|-----------------|
| 18 | COL16A1 | 7.50E-06 | 4.24E-03 | 0.18 | 1.19(1.1-1.29) |
| 19 | NUAK1 | 7.67E-06 | 4.24E-03 | 0.15 | 1.16(1.09-1.23) |
| 20 | JAM2 | 8.50E-06 | 4.24E-03 | 0.20 | 1.22(1.12-1.33) |
| 21 | GPR124 | 8.82E-06 | 4.24E-03 | 0.27 | 1.31(1.16-1.47) |
| 22 | ARNT | 9.44E-06 | 4.24E-03 | 0.33 | 1.39(1.2-1.62) |
| 23 | GALNT10 | 9.62E-06 | 4.24E-03 | 0.23 | 1.26(1.14-1.39) |
| 24 | MINOS1-NBL1 | 1.02E-05 | 4.24E-03 | 0.17 | 1.19(1.1-1.29) |
| 25 | AKAP12 | 1.11E-05 | 4.24E-03 | 0.13 | 1.14(1.08-1.21) |
| 26 | PDGFD | 1.16E-05 | 4.24E-03 | 0.15 | 1.16(1.08-1.24) |
| 27 | LRRC17 | 1.16E-05 | 4.47E-03 | 0.13 | 1.14(1.07-1.2) |
| 28 | PDGFRB | 1.29E-05 | 4.47E-03 | 0.25 | 1.28(1.14-1.43) |
| 29 | GLT8D2 | 1.30E-05 | 4.47E-03 | 0.18 | 1.2(1.1-1.3) |
| 30 | PTPRD | 1.38E-05 | 4.47E-03 | 0.20 | 1.22(1.12-1.34) |
| 31 | ZEB1 | 1.54E-05 | 4.47E-03 | 0.16 | 1.17(1.09-1.26) |
| 32 | ID3 | 1.56E-05 | 4.47E-03 | 0.17 | 1.19(1.1-1.29) |
| 33 | PDPN | 1.71E-05 | 4.47E-03 | 0.16 | 1.18(1.09-1.27) |
| 34 | VAT1 | 1.77E-05 | 4.47E-03 | 0.32 | 1.38(1.19-1.59) |
| 35 | SV2A | 1.80E-05 | 4.47E-03 | 0.23 | 1.26(1.13-1.4) |
| 36 | C1QTNF3 | 1.81E-05 | 4.47E-03 | 0.15 | 1.16(1.09-1.24) |
| 37 | MXRA8 | 1.86E-05 | 4.47E-03 | 0.16 | 1.18(1.09-1.27) |
| 38 | ZCCHC24 | 1.93E-05 | 4.47E-03 | 0.21 | 1.23(1.12-1.36) |
| 39 | DCHS1 | 1.98E-05 | 4.47E-03 | 0.18 | 1.2(1.1-1.31) |
| 40 | TUBB6 | 2.02E-05 | 4.47E-03 | 0.17 | 1.19(1.1-1.28) |
| 41 | COPZ2 | 2.05E-05 | 4.47E-03 | 0.21 | 1.24(1.12-1.37) |
| 42 | TIMP3 | 2.05E-05 | 4.47E-03 | 0.12 | 1.13(1.07-1.19) |
| 43 | FZD1 | 2.08E-05 | 4.47E-03 | 0.20 | 1.22(1.11-1.34) |
| 44 | GFPT2 | 2.09E-05 | 4.47E-03 | 0.22 | 1.24(1.12-1.37) |
| 45 | SYDE1 | 2.14E-05 | 4.47E-03 | 0.37 | 1.44(1.22-1.71) |
| 46 | HSD17B6 | 2.24E-05 | 6.38E-03 | 0.22 | 1.25(1.13-1.39) |
| 47 | SH3PXD2A | 2.29E-05 | 8.43E-03 | 0.30 | 1.35(1.18-1.56) |
| 48 | MPRIP | 2.66E-05 | 8.43E-03 | 0.32 | 1.37(1.18-1.59) |
| 49 | IGF2 | 2.75E-05 | 8.43E-03 | 0.06 | 1.06(1.03-1.09) |
| 50 | RHOBTB3 | 2.92E-05 | 8.43E-03 | 0.14 | 1.15(1.08-1.22) |
| 51 | ECM2 | 2.92E-05 | 1.24E-02 | 0.15 | 1.16(1.08-1.24) |
| 52 | MAGEL2 | 3.05E-05 | 1.78E-02 | 0.31 | 1.36(1.18-1.57) |
| 53 | HSPB6 | 3.42E-05 | 1.78E-02 | 0.23 | 1.26(1.13-1.41) |
| 54 | ENOX1 | 3.93E-05 | 1.78E-02 | 0.32 | 1.38(1.18-1.61) |
| 55 | CD302 | 4.17E-05 | 1.78E-02 | 0.17 | 1.18(1.09-1.28) |
| 56 | N4BP2L2 | 5.02E-05 | 1.78E-02 | 0.24 | 1.27(1.13-1.42) |
| 57 | FXYD5 | 5.73E-05 | 1.78E-02 | 0.18 | 1.2(1.1-1.31) |
| 58 | FOXN3 | 6.04E-05 | 1.78E-02 | 0.25 | 1.28(1.14-1.45) |
| 59 | COL5A1 | 6.06E-05 | 1.78E-02 | 0.10 | 1.1(1.05-1.16) |
| 60 | KIF26B | 6.18E-05 | 1.78E-02 | 0.20 | 1.22(1.11-1.35) |
| 61 | ITSN1 | 6.36E-05 | 1.78E-02 | 0.32 | 1.37(1.18-1.6) |

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|-----|----------|----------|----------|------|-----------------|
| 62 | SORBS1 | 6.36E-05 | 1.78E-02 | 0.21 | 1.24(1.11-1.37) |
| 63 | FGF1 | 6.46E-05 | 1.78E-02 | 0.29 | 1.33(1.16-1.54) |
| 64 | SLIT3 | 6.49E-05 | 1.78E-02 | 0.21 | 1.23(1.11-1.36) |
| 65 | AP3S1 | 6.81E-05 | 2.15E-02 | 0.30 | 1.35(1.16-1.56) |
| 66 | HIGD1B | 7.58E-05 | 2.15E-02 | 0.47 | 1.6(1.27-2.01) |
| 67 | APOD | 7.86E-05 | 2.15E-02 | 0.17 | 1.18(1.09-1.28) |
| 68 | TMEM158 | 8.08E-05 | 2.15E-02 | 0.11 | 1.12(1.06-1.18) |
| 69 | MUL1 | 8.31E-05 | 2.15E-02 | 0.44 | 1.55(1.25-1.93) |
| 70 | LEFTY2 | 8.38E-05 | 2.15E-02 | 0.23 | 1.26(1.12-1.41) |
| 71 | SACS | 8.97E-05 | 2.15E-02 | 0.21 | 1.24(1.11-1.38) |
| 72 | FSTL3 | 9.02E-05 | 2.15E-02 | 0.36 | 1.44(1.2-1.72) |
| 73 | PPP1R13L | 1.01E-04 | 2.15E-02 | 0.23 | 1.26(1.12-1.41) |
| 74 | WBP1L | 1.10E-04 | 2.15E-02 | 0.25 | 1.28(1.13-1.45) |
| 75 | ANGPTL2 | 1.12E-04 | 2.15E-02 | 0.19 | 1.21(1.1-1.33) |
| 76 | CLIP3 | 1.13E-04 | 2.15E-02 | 0.18 | 1.2(1.09-1.31) |
| 77 | CYTH3 | 1.14E-04 | 2.15E-02 | 0.36 | 1.44(1.19-1.73) |
| 78 | ESD | 1.20E-04 | 2.15E-02 | 0.25 | 1.28(1.13-1.45) |
| 79 | PLXDC1 | 1.21E-04 | 2.15E-02 | 0.20 | 1.22(1.1-1.36) |
| 80 | CYBRD1 | 1.27E-04 | 2.15E-02 | 0.13 | 1.14(1.07-1.22) |
| 81 | LMCD1 | 1.38E-04 | 2.15E-02 | 0.18 | 1.2(1.09-1.32) |
| 82 | PAK4 | 1.40E-04 | 2.15E-02 | 0.26 | 1.29(1.13-1.47) |
| 83 | SEMA4F | 1.42E-04 | 2.15E-02 | 0.34 | 1.41(1.18-1.68) |
| 84 | PCOLCE | 1.43E-04 | 2.15E-02 | 0.14 | 1.15(1.07-1.24) |
| 85 | SVEP1 | 1.43E-04 | 2.15E-02 | 0.19 | 1.21(1.1-1.33) |
| 86 | ZFP36 | 1.46E-04 | 2.15E-02 | 0.14 | 1.15(1.07-1.24) |
| 87 | ABCD2 | 1.48E-04 | 2.15E-02 | 1.06 | 2.89(1.67-4.99) |
| 88 | PPP3CA | 1.55E-04 | 2.15E-02 | 0.21 | 1.23(1.11-1.37) |
| 89 | MAN2A1 | 1.57E-04 | 2.15E-02 | 0.20 | 1.22(1.1-1.34) |
| 90 | COMP | 1.70E-04 | 2.36E-02 | 0.08 | 1.09(1.04-1.13) |
| 91 | RHOT1 | 1.74E-04 | 2.36E-02 | 0.28 | 1.32(1.14-1.53) |
| 92 | ARHGAP19 | 1.74E-04 | 2.36E-02 | 0.28 | 1.33(1.14-1.53) |
| 93 | FBN1 | 1.80E-04 | 2.36E-02 | 0.10 | 1.11(1.05-1.17) |
| 94 | PSMC4 | 1.82E-04 | 2.36E-02 | 0.20 | 1.22(1.1-1.36) |
| 95 | PRRX1 | 1.89E-04 | 2.36E-02 | 0.13 | 1.14(1.06-1.22) |
| 96 | LBP | 1.98E-04 | 2.36E-02 | 0.36 | 1.43(1.18-1.72) |
| 97 | AEBP1 | 1.99E-04 | 2.36E-02 | 0.11 | 1.11(1.05-1.17) |
| 98 | XYLT1 | 2.00E-04 | 2.36E-02 | 0.20 | 1.22(1.1-1.35) |
| 99 | PADI2 | 2.00E-04 | 2.36E-02 | 0.18 | 1.19(1.09-1.31) |
| 100 | SFXN3 | 2.01E-04 | 2.36E-02 | 0.25 | 1.28(1.13-1.46) |
| 101 | INHBA | 2.07E-04 | 2.36E-02 | 0.08 | 1.09(1.04-1.14) |
| 102 | EYA4 | 2.17E-04 | 2.36E-02 | 0.12 | 1.13(1.06-1.2) |
| 103 | PDGFRA | 2.27E-04 | 2.36E-02 | 0.10 | 1.1(1.05-1.16) |
| 104 | PTH1R | 2.31E-04 | 2.36E-02 | 0.41 | 1.5(1.21-1.86) |
| 105 | LTBP1 | 2.33E-04 | 2.36E-02 | 0.13 | 1.14(1.06-1.23) |

| | | | | | |
|-----|----------|----------|----------|------|-----------------|
| 106 | PCDH9 | 2.36E-04 | 2.36E-02 | 0.19 | 1.21(1.09-1.34) |
| 107 | TUBB2A | 2.40E-04 | 2.36E-02 | 0.12 | 1.13(1.06-1.2) |
| 108 | PALLD | 2.40E-04 | 2.36E-02 | 0.14 | 1.15(1.07-1.23) |
| 109 | AHDC1 | 2.40E-04 | 2.36E-02 | 0.19 | 1.21(1.09-1.34) |
| 110 | WASF2 | 2.41E-04 | 2.36E-02 | 0.22 | 1.25(1.11-1.41) |
| 111 | PPAP2A | 2.41E-04 | 2.36E-02 | 0.20 | 1.22(1.1-1.36) |
| 112 | FAP | 2.50E-04 | 2.36E-02 | 0.10 | 1.1(1.05-1.16) |
| 113 | APH1B | 2.51E-04 | 2.36E-02 | 0.25 | 1.29(1.12-1.47) |
| 114 | ZBTB40 | 2.53E-04 | 2.36E-02 | 0.36 | 1.44(1.18-1.74) |
| 115 | AKR7A2 | 2.59E-04 | 2.36E-02 | 0.23 | 1.26(1.11-1.43) |
| 116 | STK39 | 2.59E-04 | 3.17E-02 | 0.18 | 1.19(1.09-1.31) |
| 117 | ITGA5 | 2.71E-04 | 3.17E-02 | 0.22 | 1.25(1.11-1.41) |
| 118 | NET1 | 2.72E-04 | 3.17E-02 | 0.16 | 1.17(1.07-1.27) |
| 119 | TRIL | 2.80E-04 | 3.17E-02 | 0.18 | 1.2(1.09-1.32) |
| 120 | RBMS1P1 | 2.89E-04 | 3.17E-02 | 0.24 | 1.27(1.11-1.44) |
| 121 | SNAI2 | 2.96E-04 | 3.17E-02 | 0.11 | 1.11(1.05-1.18) |
| 122 | RCN3 | 3.05E-04 | 3.17E-02 | 0.17 | 1.19(1.08-1.31) |
| 123 | GUCY1A2 | 3.07E-04 | 3.17E-02 | 0.15 | 1.16(1.07-1.26) |
| 124 | PER1 | 3.10E-04 | 3.17E-02 | 0.37 | 1.44(1.18-1.76) |
| 125 | HTRA1 | 3.26E-04 | 3.17E-02 | 0.14 | 1.15(1.06-1.23) |
| 126 | IGFBP4 | 3.26E-04 | 3.17E-02 | 0.13 | 1.14(1.06-1.23) |
| 127 | CXCL14 | 3.40E-04 | 3.17E-02 | 0.08 | 1.08(1.04-1.13) |
| 128 | LYPD3 | 3.40E-04 | 3.17E-02 | 0.22 | 1.24(1.1-1.4) |
| 129 | FBXO17 | 3.48E-04 | 3.17E-02 | 0.15 | 1.16(1.07-1.26) |
| 130 | LHFP | 3.54E-04 | 3.17E-02 | 0.13 | 1.14(1.06-1.22) |
| 131 | LRRC15 | 3.61E-04 | 3.17E-02 | 0.10 | 1.1(1.04-1.16) |
| 132 | MPZ | 3.63E-04 | 3.17E-02 | 0.32 | 1.38(1.16-1.64) |
| 133 | DACT1 | 3.65E-04 | 3.17E-02 | 0.15 | 1.16(1.07-1.27) |
| 134 | KIAA1462 | 3.65E-04 | 3.17E-02 | 0.19 | 1.2(1.09-1.33) |
| 135 | OMD | 3.70E-04 | 3.55E-02 | 0.16 | 1.17(1.07-1.28) |
| 136 | PDE2A | 3.92E-04 | 3.55E-02 | 0.33 | 1.4(1.16-1.68) |
| 137 | RAI14 | 4.00E-04 | 3.55E-02 | 0.16 | 1.17(1.07-1.27) |
| 138 | RECK | 4.01E-04 | 3.55E-02 | 0.18 | 1.2(1.08-1.32) |
| 139 | GNB1 | 4.14E-04 | 3.55E-02 | 0.25 | 1.28(1.12-1.47) |
| 140 | TNFAIP6 | 4.14E-04 | 3.55E-02 | 0.12 | 1.13(1.06-1.21) |
| 141 | COL8A1 | 4.16E-04 | 3.55E-02 | 0.12 | 1.13(1.05-1.2) |
| 142 | RGS16 | 4.23E-04 | 3.55E-02 | 0.15 | 1.17(1.07-1.27) |
| 143 | RIN2 | 4.26E-04 | 3.55E-02 | 0.21 | 1.24(1.1-1.39) |
| 144 | ACTR1A | 4.26E-04 | 3.55E-02 | 0.25 | 1.28(1.12-1.48) |
| 145 | FMO2 | 4.32E-04 | 3.55E-02 | 0.12 | 1.13(1.06-1.21) |
| 146 | NR2F1 | 4.35E-04 | 3.55E-02 | 0.12 | 1.13(1.06-1.21) |
| 147 | DKK2 | 4.40E-04 | 3.55E-02 | 0.16 | 1.18(1.08-1.29) |
| 148 | APBB2 | 4.45E-04 | 3.55E-02 | 0.19 | 1.21(1.09-1.35) |
| 149 | OLFML1 | 4.49E-04 | 4.05E-02 | 0.16 | 1.17(1.07-1.28) |

| | | | | | |
|-----|----------|----------|----------|------|-----------------|
| 150 | SASH1 | 4.71E-04 | 4.05E-02 | 0.22 | 1.25(1.1-1.41) |
| 151 | BLMH | 4.80E-04 | 4.05E-02 | 0.22 | 1.25(1.1-1.41) |
| 152 | CPEB1 | 5.01E-04 | 4.05E-02 | 0.26 | 1.29(1.12-1.5) |
| 153 | RPS6KA2 | 5.04E-04 | 4.05E-02 | 0.24 | 1.28(1.11-1.47) |
| 154 | CALCOCO1 | 5.04E-04 | 4.05E-02 | 0.29 | 1.34(1.13-1.57) |
| 155 | PINK1 | 5.24E-04 | 4.05E-02 | 0.28 | 1.32(1.13-1.54) |
| 156 | ANTXR1 | 5.25E-04 | 4.05E-02 | 0.18 | 1.2(1.08-1.33) |
| 157 | RPS23P8 | 5.37E-04 | 4.05E-02 | 0.31 | 1.37(1.14-1.63) |
| 158 | ADH1B | 5.50E-04 | 4.05E-02 | 0.08 | 1.09(1.04-1.14) |
| 159 | GPC1 | 5.53E-04 | 4.05E-02 | 0.19 | 1.21(1.09-1.35) |
| 160 | PCBP3 | 5.56E-04 | 4.05E-02 | 0.44 | 1.56(1.21-2.01) |
| 161 | TMEM45A | 5.57E-04 | 4.05E-02 | 0.11 | 1.12(1.05-1.19) |
| 162 | DCN | 5.65E-04 | 4.05E-02 | 0.10 | 1.1(1.04-1.16) |
| 163 | PPP1R3C | 5.69E-04 | 4.05E-02 | 0.14 | 1.15(1.06-1.24) |
| 164 | RARRES2 | 5.73E-04 | 4.05E-02 | 0.12 | 1.12(1.05-1.2) |
| 165 | PKD2 | 5.77E-04 | 4.05E-02 | 0.20 | 1.22(1.09-1.36) |
| 166 | MFAP5 | 5.85E-04 | 4.05E-02 | 0.07 | 1.08(1.03-1.12) |
| 167 | PLOD3 | 5.85E-04 | 4.05E-02 | 0.30 | 1.34(1.14-1.59) |
| 168 | SPARC | 5.87E-04 | 4.05E-02 | 0.13 | 1.14(1.06-1.23) |
| 169 | RHOB | 5.92E-04 | 4.05E-02 | 0.13 | 1.14(1.06-1.22) |
| 170 | OSR2 | 5.95E-04 | 4.05E-02 | 0.11 | 1.11(1.05-1.19) |
| 171 | SUPT5H | 5.98E-04 | 4.05E-02 | 0.22 | 1.25(1.1-1.41) |
| 172 | PDP1 | 6.01E-04 | 4.05E-02 | 0.16 | 1.17(1.07-1.28) |
| 173 | TPT1 | 6.23E-04 | 4.05E-02 | 0.39 | 1.48(1.18-1.85) |
| 174 | SNORA31 | 6.32E-04 | 4.05E-02 | 0.40 | 1.49(1.19-1.88) |
| 175 | PDLIM2 | 6.37E-04 | 4.05E-02 | 0.18 | 1.19(1.08-1.32) |
| 176 | MOXD1 | 6.39E-04 | 4.05E-02 | 0.14 | 1.15(1.06-1.25) |
| 177 | THBS2 | 6.46E-04 | 4.05E-02 | 0.07 | 1.08(1.03-1.12) |
| 178 | APPL2 | 6.53E-04 | 4.05E-02 | 0.19 | 1.21(1.08-1.35) |
| 179 | ADRA2A | 6.55E-04 | 4.05E-02 | 0.12 | 1.13(1.05-1.21) |
| 180 | LMOD1 | 6.56E-04 | 4.05E-02 | 0.18 | 1.19(1.08-1.32) |
| 181 | RPL21P75 | 6.57E-04 | 4.05E-02 | 0.27 | 1.31(1.12-1.53) |
| 182 | TGFB1I1 | 6.69E-04 | 4.05E-02 | 0.12 | 1.12(1.05-1.2) |
| 183 | YWHAB | 6.75E-04 | 4.05E-02 | 0.27 | 1.31(1.12-1.53) |
| 184 | GNAI1 | 6.80E-04 | 4.05E-02 | 0.16 | 1.17(1.07-1.28) |
| 185 | TMEM222 | 6.81E-04 | 4.05E-02 | 0.35 | 1.42(1.16-1.73) |
| 186 | SERTAD3 | 6.92E-04 | 4.05E-02 | 0.18 | 1.2(1.08-1.33) |
| 187 | SNORA21 | 6.97E-04 | 4.05E-02 | 0.27 | 1.31(1.12-1.53) |
| 188 | PRKG1 | 6.98E-04 | 4.05E-02 | 0.25 | 1.28(1.11-1.48) |
| 189 | SRPX2 | 7.07E-04 | 4.05E-02 | 0.16 | 1.17(1.07-1.28) |
| 190 | UBL3 | 7.15E-04 | 4.05E-02 | 0.22 | 1.24(1.1-1.41) |
| 191 | NDRG3 | 7.21E-04 | 4.05E-02 | 0.24 | 1.27(1.1-1.45) |
| 192 | ZNF365 | 7.27E-04 | 4.05E-02 | 0.33 | 1.39(1.15-1.67) |
| 193 | CPQ | 7.30E-04 | 4.05E-02 | 0.20 | 1.22(1.09-1.38) |

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|-----|----------------|----------|----------|------|-----------------|
| 194 | RNASEL | 7.34E-04 | 4.05E-02 | 0.25 | 1.29(1.11-1.49) |
| 195 | PLEKHA1 | 7.45E-04 | 4.05E-02 | 0.21 | 1.24(1.09-1.4) |
| 196 | HIPK1 | 7.46E-04 | 4.05E-02 | 0.14 | 1.15(1.06-1.24) |
| 197 | PTRF | 7.51E-04 | 4.05E-02 | 0.15 | 1.17(1.07-1.28) |
| 198 | FLNC | 7.53E-04 | 4.05E-02 | 0.22 | 1.25(1.1-1.42) |
| 199 | PDLIM3 | 7.54E-04 | 4.05E-02 | 0.11 | 1.12(1.05-1.2) |
| 200 | ITPKC | 7.54E-04 | 4.05E-02 | 0.22 | 1.25(1.1-1.42) |
| 201 | HSPB2-C11orf52 | 7.55E-04 | 4.05E-02 | 0.17 | 1.19(1.08-1.31) |
| 202 | ASPN | 7.56E-04 | 4.05E-02 | 0.08 | 1.08(1.03-1.13) |
| 203 | NPY | 7.62E-04 | 4.05E-02 | 0.11 | 1.11(1.05-1.18) |
| 204 | KIAA0355 | 7.64E-04 | 4.05E-02 | 0.23 | 1.26(1.1-1.44) |
| 205 | KPNA3 | 7.75E-04 | 4.05E-02 | 0.24 | 1.27(1.1-1.45) |
| 206 | BCHE | 7.88E-04 | 4.37E-02 | 0.12 | 1.13(1.05-1.21) |
| 207 | SUGCT | 7.97E-04 | 4.37E-02 | 0.14 | 1.15(1.06-1.25) |
| 208 | PPEF1 | 8.15E-04 | 4.37E-02 | 0.30 | 1.35(1.13-1.61) |
| 209 | ZNF780B | 8.28E-04 | 4.37E-02 | 0.29 | 1.34(1.13-1.58) |
| 210 | RPS16 | 8.42E-04 | 4.37E-02 | 0.32 | 1.38(1.14-1.67) |
| 211 | CRY2 | 8.52E-04 | 4.37E-02 | 0.38 | 1.47(1.17-1.83) |
| 212 | ERCC8 | 8.67E-04 | 4.37E-02 | 0.25 | 1.29(1.11-1.49) |
| 213 | MYL9 | 8.85E-04 | 4.37E-02 | 0.12 | 1.13(1.05-1.21) |
| 214 | COL10A1 | 8.88E-04 | 4.37E-02 | 0.08 | 1.09(1.03-1.14) |
| 215 | CLIC3 | 8.92E-04 | 4.37E-02 | 0.11 | 1.12(1.05-1.19) |
| 216 | PTPN1 | 9.00E-04 | 4.37E-02 | 0.30 | 1.35(1.13-1.61) |
| 217 | EMILIN1 | 9.11E-04 | 4.37E-02 | 0.16 | 1.18(1.07-1.3) |
| 218 | COX7C | 9.22E-04 | 4.37E-02 | 0.25 | 1.28(1.11-1.49) |
| 219 | TMEM47 | 9.28E-04 | 4.37E-02 | 0.09 | 1.1(1.04-1.16) |
| 220 | FAM114A1 | 9.37E-04 | 4.37E-02 | 0.22 | 1.24(1.09-1.41) |
| 221 | EMCN | 9.42E-04 | 4.37E-02 | 0.22 | 1.25(1.09-1.42) |
| 222 | EDNRA | 9.42E-04 | 4.37E-02 | 0.12 | 1.13(1.05-1.21) |
| 223 | COL11A1 | 9.44E-04 | 4.37E-02 | 0.05 | 1.05(1.02-1.09) |
| 224 | ACTN4 | 9.50E-04 | 4.37E-02 | 0.18 | 1.2(1.08-1.33) |
| 225 | GFRA1 | 9.61E-04 | 4.37E-02 | 0.20 | 1.23(1.09-1.38) |
| 226 | SARS2 | 9.81E-04 | 4.37E-02 | 0.18 | 1.19(1.07-1.32) |
| 227 | COX7CP1 | 9.81E-04 | 4.37E-02 | 0.24 | 1.27(1.1-1.47) |
| 228 | ENPP1 | 9.82E-04 | 4.37E-02 | 0.15 | 1.16(1.06-1.27) |
| 229 | CRISPLD2 | 9.83E-04 | 4.37E-02 | 0.09 | 1.1(1.04-1.16) |
| 230 | CDC42 | 9.86E-04 | 4.37E-02 | 0.16 | 1.17(1.07-1.29) |
| 231 | ACTA2 | 9.89E-04 | 4.37E-02 | 0.09 | 1.1(1.04-1.16) |

Abbreviations: pos positively associated, neg negatively associated, P P value, FDR faluse discovery rate, C Coefficient, HR Hazard Ratio, CI confidence interval.

Supplementary Table S2. The loadings of 341 genes in the first 2 components.

| Genes | PC1 | PC2 | Genes | PC1 | PC2 | Genes | PC1 | PC2 |
|----------|--------|--------|--------------------|--------|--------|-----------|--------|--------|
| AADAC | 0.008 | -0.059 | GJB1 | 0.022 | -0.059 | PPP3CA | -0.026 | 0.014 |
| ABCD2 | 0.000 | 0.002 | GLT8D2 | -0.089 | 0.020 | PRIM2 | 0.011 | -0.010 |
| ACOT13 | 0.010 | -0.052 | GMPR | 0.037 | -0.091 | PRKG1 | -0.025 | 0.014 |
| ACTA2 | -0.137 | -0.049 | GMPR2 | 0.005 | -0.010 | PRPS2 | 0.010 | -0.062 |
| ACTN4 | -0.016 | 0.014 | GNAI1 | -0.015 | 0.070 | PRRX1 | -0.092 | -0.019 |
| ACTR1A | -0.019 | 0.005 | GNB1 | -0.011 | 0.013 | PSMC4 | -0.006 | 0.007 |
| ADH1B | -0.108 | -0.016 | GPC1 | -0.027 | 0.073 | PSME2P2 | 0.011 | -0.107 |
| ADRA2A | -0.060 | 0.042 | GPR124 | -0.048 | 0.032 | PTH1R | -0.005 | 0.026 |
| AEBP1 | -0.149 | -0.012 | GPRC5C | 0.012 | -0.032 | PTPN1 | -0.010 | 0.002 |
| AHDC1 | 0.001 | 0.058 | GSTZ1 | 0.014 | -0.045 | PTPRD | -0.038 | 0.023 |
| AKAP12 | -0.056 | 0.076 | GUCY1A2 | -0.016 | 0.024 | PTRF | -0.082 | 0.032 |
| AKR7A2 | 0.000 | 0.009 | GZMB | -0.012 | -0.124 | RAB11FIP5 | -0.016 | 0.027 |
| ALDH5A1 | 0.035 | -0.037 | HIGD1B | -0.008 | 0.004 | RAI14 | -0.048 | 0.030 |
| ANGPTL2 | -0.076 | 0.013 | HIPK1 | -0.058 | 0.007 | RARRES2 | -0.081 | 0.003 |
| ANTXR1 | -0.049 | 0.005 | HLA.DOB | 0.013 | -0.084 | RBMS1P1 | -0.016 | 0.033 |
| AP3S1 | -0.018 | 0.000 | HP1BP3 | 0.000 | 0.012 | RCL1 | 0.010 | -0.023 |
| APBB2 | -0.036 | 0.043 | HSD17B6 | -0.055 | -0.012 | RCN3 | -0.071 | 0.005 |
| APH1B | 0.005 | 0.005 | HSPB2.C11or f52 | -0.017 | 0.033 | RECK | -0.063 | 0.015 |
| APOD | -0.045 | 0.038 | HSPB6 | -0.004 | 0.059 | RGS16 | -0.066 | -0.020 |
| APPL2 | -0.005 | 0.052 | HSPB7 | -0.015 | 0.017 | RHOB | -0.042 | 0.042 |
| ARHGAP19 | 0.002 | 0.018 | HTRA1 | -0.083 | 0.015 | RHOBTB3 | -0.075 | 0.103 |
| ARNT | -0.002 | 0.028 | ID3 | -0.074 | 0.027 | RHOT1 | -0.004 | 0.028 |
| ASAP3 | -0.014 | 0.048 | IGF2 | -0.094 | 0.387 | RIN2 | -0.033 | 0.011 |
| ASPN | -0.167 | -0.080 | IGFBP4 | -0.086 | 0.039 | RMI1 | 0.018 | -0.058 |
| AXL | -0.047 | 0.012 | IL6R | 0.018 | -0.098 | RNASEL | -0.010 | 0.004 |
| BCAP31 | 0.006 | -0.057 | INHBA | -0.178 | -0.110 | RPL21P75 | -0.001 | 0.011 |
| BCHE | -0.064 | 0.005 | ITGA5 | -0.055 | -0.025 | RPS16 | 0.001 | 0.013 |
| BLMH | -0.013 | 0.039 | ITPKC | 0.001 | 0.024 | RPS23P8 | 0.000 | 0.026 |
| BRCC3 | 0.016 | -0.068 | ITSN1 | -0.016 | 0.042 | RPS6KA2 | -0.008 | 0.026 |
| C1QTNF3 | -0.083 | 0.001 | JAM2 | -0.048 | 0.027 | SACS | -0.018 | 0.024 |
| C6orf165 | 0.007 | -0.002 | KIAA0020 | 0.008 | -0.046 | SARS2 | 0.002 | 0.049 |
| C9orf116 | 0.019 | -0.005 | KIAA0355 | -0.016 | 0.037 | SASH1 | -0.019 | 0.027 |
| CAAP1 | 0.014 | -0.043 | KIAA1462 | -0.030 | 0.040 | SDF2L1 | -0.001 | -0.065 |
| CALCOCO1 | -0.006 | 0.033 | KIF26B | -0.048 | 0.006 | SDHAP3 | 0.011 | -0.021 |
| CASP8AP2 | 0.006 | -0.013 | KPNA3 | -0.015 | 0.013 | SDR39U1 | 0.009 | 0.006 |
| CCL13 | 0.000 | -0.027 | KRT18P38 | 0.004 | -0.002 | SEMA4F | 0.002 | 0.034 |
| CD302 | -0.052 | -0.001 | LBP | -0.001 | 0.008 | SENP6 | -0.001 | 0.002 |

| CD38 | 0.003 | -0.099 | LEFTY2 | -0.005 | 0.058 | SERTAD3 | -0.010 | 0.025 | |
|----------|--------|--------|-----------------|--------|--------|----------|--------|--------|-------|
| CDC42 | -0.012 | -0.015 | LHFP | -0.100 | 0.001 | SFXN3 | -0.035 | 0.004 | |
| CHAF1B | 0.012 | -0.030 | LINC00338 | 0.017 | -0.012 | SH3PXD2A | -0.029 | 0.017 | |
| CIB1 | 0.013 | -0.069 | LMCD1 | -0.036 | -0.014 | SIRT5 | 0.015 | -0.038 | |
| CILP | -0.095 | -0.001 | LMOD1 | -0.056 | 0.024 | SLC25A1 | 0.003 | -0.040 | |
| CLIC3 | -0.050 | 0.031 | LPAR2 | 0.014 | 0.009 | SLC27A2 | 0.013 | -0.025 | |
| CLIP3 | -0.025 | 0.089 | LRIF1 | 0.007 | -0.009 | SLC37A4 | 0.017 | -0.006 | |
| CNOT4 | 0.007 | -0.023 | LRIG1 | 0.018 | -0.011 | SLC5A1 | 0.023 | -0.085 | |
| COA4 | 0.015 | -0.058 | LRP1 | -0.038 | 0.037 | SLC7A11 | 0.019 | -0.070 | |
| COL10A1 | -0.153 | -0.076 | LRRC15 | -0.130 | -0.068 | SLIT3 | -0.023 | 0.073 | |
| COL11A1 | -0.237 | -0.152 | LRRC17 | -0.075 | 0.067 | SNAI2 | -0.133 | -0.034 | |
| COL16A1 | -0.088 | 0.014 | LRRK1 | 0.002 | -0.020 | SNORA21 | -0.002 | 0.039 | |
| COL5A1 | -0.169 | -0.038 | LRRN3 | -0.008 | 0.023 | SNORA31 | -0.009 | 0.013 | |
| COL8A1 | -0.101 | -0.029 | LTBP1 | -0.028 | 0.060 | SNRPA1 | 0.014 | -0.056 | |
| COMP | -0.132 | -0.028 | LUZP1 | -0.013 | 0.022 | SORBS1 | -0.028 | 0.015 | |
| COPZ2 | -0.072 | -0.013 | LYPD3 | 0.004 | 0.027 | SPARC | -0.100 | -0.025 | |
| COX7A2P2 | 0.011 | -0.038 | MAGEF1 | 0.007 | 0.001 | SPDEF | 0.041 | -0.033 | |
| COX7C | 0.005 | 0.013 | MAGEL2 | -0.012 | 0.051 | SRPX2 | -0.074 | -0.011 | |
| COX7CP1 | 0.005 | 0.011 | MAK | 0.024 | 0.004 | ST6GAL1 | 0.010 | -0.056 | |
| CPEB1 | -0.004 | 0.036 | MAL | -0.026 | 0.081 | STK39 | -0.016 | 0.007 | |
| CPQ | -0.031 | 0.005 | MAN2A1 | -0.030 | -0.045 | STOML2 | 0.016 | -0.041 | |
| CREB3 | -0.001 | -0.037 | MCAT | 0.011 | -0.025 | STXBP6 | 0.032 | -0.123 | |
| CRISPLD2 | -0.145 | -0.049 | MCM3 | 0.017 | -0.016 | SUGCT | -0.070 | -0.018 | |
| CRY2 | -0.004 | 0.017 | MCM5 | 0.008 | -0.059 | SUPT5H | -0.006 | 0.025 | |
| CRYAB | -0.054 | -0.017 | MFAP4 | -0.067 | 0.103 | SURF2 | 0.011 | -0.017 | |
| CXCL11 | -0.008 | -0.277 | MFAP5 | -0.122 | 0.009 | SV2A | -0.005 | 0.068 | |
| CXCL14 | -0.151 | -0.028 | MINOS1.NB L1 | -0.093 | -0.018 | SVEP1 | -0.065 | -0.011 | |
| CYB561 | 0.012 | -0.022 | | MOXD1 | -0.065 | -0.027 | SVIL | -0.041 | 0.045 |
| CYBA | 0.007 | -0.106 | MP RIP | -0.022 | 0.050 | SYDE1 | -0.018 | 0.035 | |
| CYBRD1 | -0.023 | 0.016 | MPZ | 0.002 | 0.019 | SYK | 0.018 | -0.040 | |
| CYTH3 | -0.010 | 0.019 | MRPS11 | 0.009 | -0.045 | SYNGR2 | 0.002 | -0.085 | |
| DACT1 | -0.076 | 0.020 | MUL1 | 0.002 | 0.003 | TAP1 | 0.006 | -0.216 | |
| DCHS1 | -0.070 | 0.073 | MXRA8 | -0.077 | 0.071 | TAP2 | 0.005 | -0.134 | |
| DCN | -0.143 | -0.036 | MYL9 | -0.066 | 0.022 | TBCC | 0.011 | -0.011 | |
| DHRS4 | 0.012 | -0.018 | N4BP2L2 | -0.004 | 0.015 | TDP2 | 0.005 | -0.045 | |
| DKK2 | -0.047 | 0.006 | NAT10 | 0.017 | -0.014 | TESK1 | 0.001 | -0.022 | |
| DNAJA1 | 0.006 | -0.066 | NDRG3 | 0.001 | 0.027 | TGFB1I1 | -0.111 | -0.005 | |
| ECM2 | -0.099 | 0.003 | NET1 | -0.029 | -0.010 | THBS2 | -0.188 | -0.137 | |
| EDNRA | -0.093 | -0.020 | NFX1 | 0.009 | -0.023 | THOP1 | 0.007 | -0.016 | |

| | | | | | | | | |
|----------|--------|--------|----------|--------|--------|--------------------|--------|--------|
| EFHC2 | 0.038 | -0.003 | NPY | 0.000 | 0.072 | TIMP3 | -0.133 | 0.017 |
| EFTUD1 | 0.007 | -0.027 | NR2F1 | -0.049 | 0.123 | TIPIN | 0.014 | -0.044 |
| EHD2 | -0.044 | 0.018 | NT5C | 0.008 | -0.011 | TMEM158 | -0.121 | -0.074 |
| EMC9 | 0.009 | -0.029 | NUAK1 | -0.111 | -0.004 | TMEM204 | -0.042 | 0.009 |
| EMCN | -0.029 | 0.001 | OLFML1 | -0.074 | 0.011 | TMEM222 | 0.004 | 0.006 |
| EMILIN1 | -0.066 | 0.015 | OMD | -0.068 | -0.013 | TMEM256. PLSCR3 | -0.025 | 0.039 |
| ENDOG | 0.013 | -0.042 | OSR2 | -0.060 | 0.043 | TMEM45A | -0.087 | 0.080 |
| ENOX1 | -0.030 | 0.016 | OXLD1 | 0.015 | -0.049 | TMEM47 | -0.069 | 0.053 |
| ENPP1 | -0.058 | 0.008 | PADI2 | -0.005 | -0.006 | TNFAIP6 | -0.092 | -0.106 |
| ERCC8 | -0.005 | 0.027 | PAK4 | 0.004 | 0.045 | TPT1 | -0.009 | 0.012 |
| ESD | -0.012 | 0.026 | PALLD | -0.098 | 0.005 | TRIL | -0.032 | 0.053 |
| EYA4 | -0.016 | 0.124 | PCBP3 | 0.001 | 0.009 | TRIM26 | 0.010 | -0.037 |
| FAH | 0.002 | -0.041 | PCDH9 | -0.010 | 0.058 | TRIM27 | 0.014 | -0.012 |
| FAM114A1 | -0.038 | -0.004 | PCK2 | 0.009 | -0.070 | TUBB2A | -0.073 | 0.020 |
| FAM86B2 | 0.011 | -0.005 | PCOLCE | -0.100 | 0.023 | TUBB6 | -0.074 | 0.002 |
| FAP | -0.151 | -0.102 | PDE2A | -0.012 | 0.023 | UBD | -0.004 | -0.282 |
| FBN1 | -0.156 | -0.043 | PDGFD | -0.081 | 0.037 | UBL3 | -0.018 | 0.026 |
| FBXO17 | -0.002 | 0.087 | PDGFRA | -0.126 | 0.020 | VAT1 | -0.026 | 0.030 |
| FCHSD2 | 0.003 | -0.031 | PDGFRB | -0.061 | 0.013 | VCP | 0.000 | -0.047 |
| FEN1 | 0.012 | -0.055 | PDLIM2 | -0.054 | -0.007 | WARS | -0.003 | -0.118 |
| FGF1 | -0.033 | -0.002 | PDLIM3 | -0.114 | -0.050 | WASF2 | -0.004 | 0.019 |
| FLNC | -0.019 | 0.047 | PDP1 | -0.012 | -0.030 | WBP1L | -0.032 | -0.008 |
| FMO2 | -0.039 | -0.035 | PDPN | -0.087 | 0.018 | WDR3 | 0.024 | -0.014 |
| FN3KRP | 0.005 | -0.023 | PER1 | -0.007 | 0.016 | WDR45B | 0.007 | -0.057 |
| FOCAD | 0.007 | -0.019 | PINK1 | -0.006 | 0.011 | WDR77 | 0.023 | -0.010 |
| FOXJ1 | 0.040 | 0.003 | PIR | 0.026 | -0.103 | XYLT1 | -0.057 | -0.002 |
| FOXN3 | -0.020 | 0.019 | PKD2 | -0.048 | 0.005 | YWHAH | -0.008 | 0.009 |
| FRYL | -0.002 | -0.032 | PLAA | 0.005 | -0.047 | ZBTB40 | 0.001 | 0.036 |
| FSTL3 | -0.024 | 0.015 | PLEKHA1 | -0.009 | 0.046 | ZCCHC24 | -0.073 | 0.018 |
| FXYD5 | -0.035 | -0.005 | PLOD3 | -0.012 | -0.002 | ZEB1 | -0.102 | 0.009 |
| FZD1 | -0.073 | -0.004 | PLXDC1 | -0.056 | -0.006 | ZFHX4 | -0.044 | 0.024 |
| GALNT10 | -0.054 | -0.007 | POLA2 | 0.013 | -0.036 | ZFP36 | -0.045 | -0.013 |
| GALNT12 | 0.021 | -0.029 | POLD3 | 0.015 | -0.025 | ZNF165 | 0.015 | -0.033 |
| GALNT6 | 0.018 | -0.066 | POLG | 0.008 | -0.025 | ZNF365 | -0.012 | 0.014 |
| GCH1 | -0.009 | -0.123 | PPAP2A | -0.054 | -0.004 | ZNF440 | 0.010 | 0.005 |
| GCNT1 | -0.001 | -0.025 | PPEF1 | -0.024 | -0.009 | ZNF780B | 0.002 | 0.019 |
| GFPT2 | -0.068 | -0.015 | PPP1R13L | -0.017 | 0.030 | ZNHIT2 | 0.015 | -0.019 |
| GFRA1 | -0.009 | 0.013 | PPP1R3C | -0.036 | 0.036 | | | |

Supplementary Table S3. The coefficients of first 2 component in principal component analysis.

| PC | Coef | SE | Z | P | HR |
|-----|----------|---------|-------|----------|---------|
| PC1 | -0.02862 | 0.00464 | -6.17 | 7.00E-10 | 0.97178 |
| PC2 | 0.06856 | 0.00964 | 7.11 | 1.10E-12 | 1.07097 |

Supplementary Table S4. List of 155 OS-associated genes with FDR<0.05 in the Riester et al. study.

| Genes | FDR | Genes | FDR | Genes | FDR | Genes | FDR |
|-----------|----------|-----------|----------|----------|----------|----------|----------|
| GMPR | 2.56E-04 | TESK1 | 2.33E-02 | LTA4H | 2.99E-02 | KIF20B | 4.19E-02 |
| MFAP4 | 7.61E-04 | MATN3 | 2.36E-02 | BMI1 | 2.99E-02 | TMEM134 | 4.19E-02 |
| LRRC17 | 7.61E-04 | ITGB7 | 2.42E-02 | RB1 | 3.05E-02 | PDGFRA | 4.19E-02 |
| GJB1 | 2.24E-03 | ISG20 | 2.45E-02 | PIR | 3.05E-02 | WDR77 | 4.19E-02 |
| NUAK1 | 2.24E-03 | ADH1B | 2.45E-02 | CHD1 | 3.05E-02 | SAT1 | 4.19E-02 |
| LEFTY2 | 2.24E-03 | HLA.DOB | 2.45E-02 | PDE1A | 3.15E-02 | ARID5B | 4.20E-02 |
| DDB2 | 2.24E-03 | EXTL2 | 2.45E-02 | APBB2 | 3.15E-02 | NT5C | 4.23E-02 |
| EDNRA | 2.29E-03 | COL16A1 | 2.49E-02 | GSTZ1 | 3.19E-02 | TAP1 | 4.23E-02 |
| HSD17B6 | 3.98E-03 | URI1 | 2.49E-02 | COL8A1 | 3.28E-02 | PDGFRB | 4.23E-02 |
| ZCCHC24 | 4.20E-03 | SLC20A2 | 2.49E-02 | AQP5 | 3.28E-02 | ITPKC | 4.24E-02 |
| AP3S1 | 5.90E-03 | JMJD1C | 2.49E-02 | RASA1 | 3.28E-02 | CLDN10 | 4.33E-02 |
| FOXJ1 | 9.64E-03 | FERMT2 | 2.49E-02 | SEMA4F | 3.28E-02 | P4HA2 | 4.33E-02 |
| RECK | 9.74E-03 | KCNV1 | 2.49E-02 | NCOA3 | 3.37E-02 | SPOCK1 | 4.33E-02 |
| PI3 | 1.26E-02 | NFS1 | 2.49E-02 | CUX1 | 3.37E-02 | MARK4 | 4.33E-02 |
| RAB11FIP5 | 1.26E-02 | PDGFD | 2.49E-02 | RECQL | 3.41E-02 | HOXA5 | 4.33E-02 |
| WDR45L | 1.37E-02 | GALNT10 | 2.49E-02 | SNAI2 | 3.75E-02 | SYDE1 | 4.54E-02 |
| SVIL | 1.37E-02 | AVPI1 | 2.49E-02 | RARB | 3.75E-02 | NR1H3 | 4.54E-02 |
| GULP1 | 1.37E-02 | DCN | 2.49E-02 | CD38 | 3.75E-02 | CDC23 | 4.54E-02 |
| APC | 1.37E-02 | GPC1 | 2.49E-02 | COL11A1 | 3.83E-02 | C6orf108 | 4.54E-02 |
| STK39 | 1.40E-02 | FSTL1 | 2.49E-02 | BCHE | 3.86E-02 | APPL2 | 4.54E-02 |
| XRCC4 | 1.44E-02 | N4BP2L2 | 2.49E-02 | DSE | 3.86E-02 | RIN2 | 4.57E-02 |
| CSNK1G3 | 1.44E-02 | NINJ1 | 2.63E-02 | PDS5B | 3.86E-02 | HSD3B2 | 4.57E-02 |
| CD302 | 1.65E-02 | CTNNBL1 | 2.63E-02 | SYT11 | 3.86E-02 | PAF1 | 4.60E-02 |
| ALDH1A2 | 1.66E-02 | RHOT1 | 2.63E-02 | SNTB2 | 3.86E-02 | UCP2 | 4.64E-02 |
| ECM2 | 1.68E-02 | APOD | 2.63E-02 | TNFAIP6 | 3.90E-02 | RPS6KA2 | 4.64E-02 |
| AKAP12 | 1.82E-02 | ST6GAL1 | 2.63E-02 | LMO7 | 4.04E-02 | CETN2 | 4.75E-02 |
| PDPN | 1.82E-02 | HIST1H2BJ | 2.64E-02 | TUFT1 | 4.10E-02 | LMOD1 | 4.76E-02 |
| ZEB1 | 1.82E-02 | SPDEF | 2.76E-02 | GRHPR | 4.12E-02 | LTBP1 | 4.76E-02 |
| SIRT5 | 1.82E-02 | CILP | 2.76E-02 | CCDC41 | 4.12E-02 | UTP14C | 4.76E-02 |
| EYA4 | 1.97E-02 | IGFBP6 | 2.76E-02 | COL5A2 | 4.12E-02 | GSTP1 | 4.76E-02 |
| SPATA2 | 1.97E-02 | PSMC4 | 2.76E-02 | CNN1 | 4.12E-02 | TJP1 | 4.76E-02 |
| DUS1L | 2.06E-02 | FBL | 2.76E-02 | GFPT2 | 4.12E-02 | ITSN1 | 4.76E-02 |
| SLC7A11 | 2.09E-02 | PALLD | 2.76E-02 | HLA.DOA | 4.12E-02 | PSMB8 | 4.84E-02 |
| KIAA0564 | 2.19E-02 | TIMP3 | 2.76E-02 | PTPRCAP | 4.12E-02 | PSMD8 | 4.86E-02 |
| ADARB1 | 2.19E-02 | PTPN1 | 2.76E-02 | GGNBP2 | 4.12E-02 | IL2RG | 4.87E-02 |
| LGALS3BP | 2.19E-02 | KRT23 | 2.77E-02 | MAN1B1 | 4.12E-02 | COMP | 4.89E-02 |
| GZMB | 2.19E-02 | ALDH5A1 | 2.78E-02 | SAMD4A | 4.12E-02 | ADNP | 4.89E-02 |
| CXCL14 | 2.32E-02 | CXCL11 | 2.90E-02 | TRIL | 4.12E-02 | NKG7 | 4.99E-02 |
| PJA2 | 2.33E-02 | FXYD1 | 2.96E-02 | PAC SIN3 | 4.19E-02 | | |

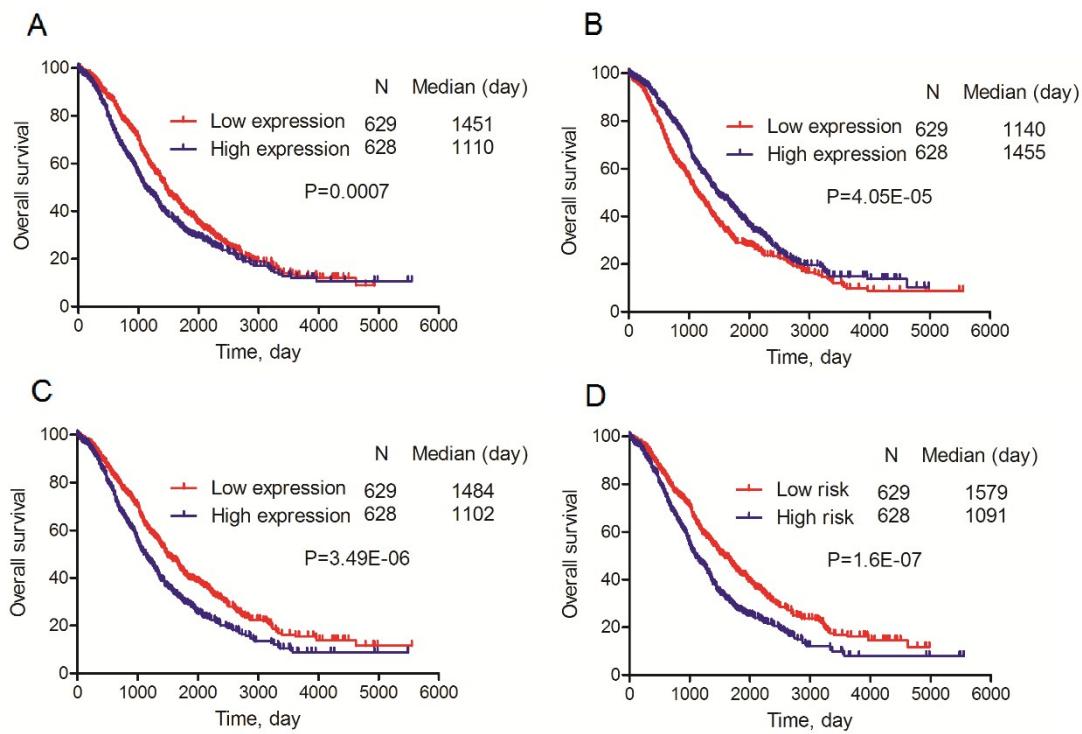
Supplementary Table S5. List of 31 OS-associated genes with FDR<0.05 in the Willis et al. study.

| Genes | FDR | Genes | FDR | Genes | FDR | Genes | FDR |
|-----------|----------|--------|----------|---------|----------|---------|----------|
| AXL | 2.20E-02 | CYBRD1 | 4.10E-02 | PAXIP1 | 4.10E-02 | ASAP3 | 4.80E-02 |
| LRIG1 | 2.20E-02 | PINK1 | 4.10E-02 | CYB561 | 4.10E-02 | FAM174B | 4.80E-02 |
| APC | 2.90E-02 | LRRN3 | 4.10E-02 | POLA2 | 4.10E-02 | AGR2 | 4.80E-02 |
| SLC33A1 | 3.00E-02 | AQP1 | 4.10E-02 | CDH1 | 4.10E-02 | SDR39U1 | 4.80E-02 |
| NUCB2 | 3.00E-02 | DES | 4.10E-02 | GMNN | 4.10E-02 | MAGT1 | 4.80E-02 |
| POLD3 | 3.00E-02 | ESR2 | 4.10E-02 | SLC37A4 | 4.10E-02 | GJB1 | 4.90E-02 |
| RAB11FIP5 | 4.10E-02 | GOLPH3 | 4.10E-02 | XRCC4 | 4.70E-02 | SDF2L1 | 5.00E-02 |
| C19orf2 | 4.10E-02 | XBP1 | 4.10E-02 | BCHE | 4.80E-02 | | |

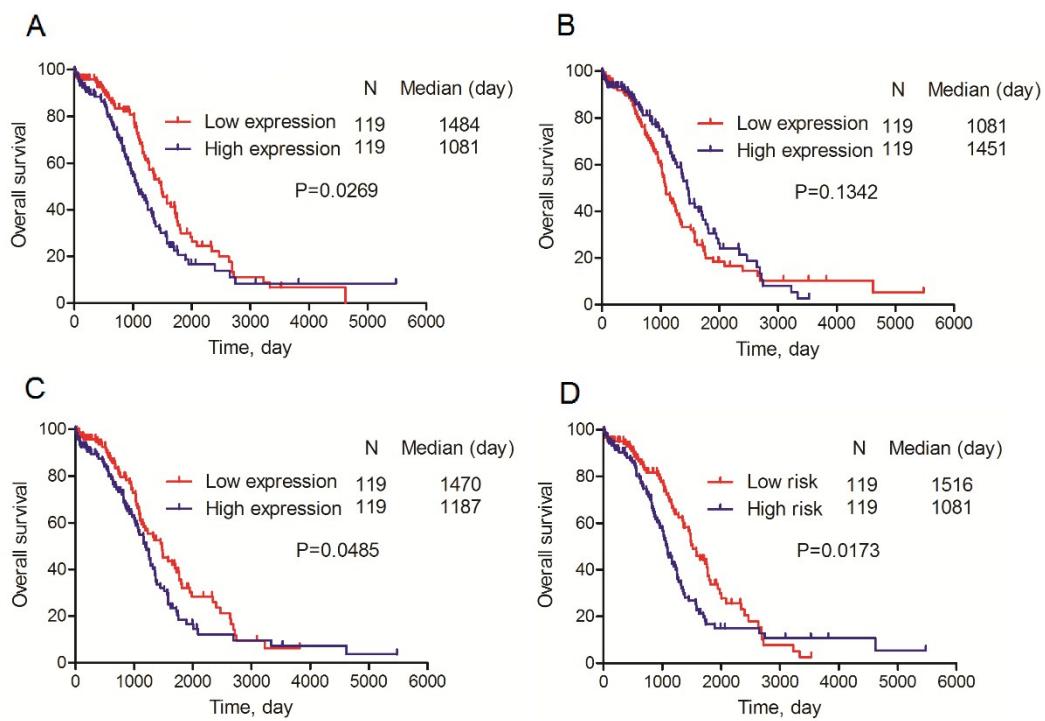
Supplementary Table S6. Univariate cox regression analysis of BCHE, GJB1 and RAB11FIP5 in EB dataset and TCGA RNA-Seq dataset.

| Variable | C | P | HR |
|---|---------|----------------|-------|
| Training in EB dataset | | | |
| BCHE | 0.1225 | 0.00079 | 1.13 |
| GJB1 | -0.2598 | 4.0E-06 | 0.77 |
| RAB11FIP5 | 0.4037 | 2.9E-06 | 1.50 |
| Combination | 2.325 | 3.6E-11 | 10.22 |
| Validation in TCGA RNA-Seq dataset | | | |
| BCHE | 0.0702 | 0.054 | 1.07 |
| GJB1 | -0.0943 | 0.026 | 0.91 |
| RAB11FIP5 | 0.237 | 0.053 | 1.27 |
| Combination | 1.000 | 0.0029 | 2.72 |

Abbreviations: C Coefficient, P P value, HR Hazard Ratio.



Supplementary Figure 1. K-M curve of 3 genes in this study which were significantly associated with overall survival in all three studies. (A) BCHE. (B) GJB1. (C) RAB11FIP5. (D) Combination of 3 genes.



Supplementary Figure 2. K-M curve of 3 genes in TCGA RNA-Seq data which were significantly associated with overall survival in all three studies. (A) BCHE. (B) GJB1. (C) RAB11FIP5. (D) Combination of 3 genes.