Identification of cancer prognosis-associated functional modules using differential co-expression networks

SUPPLEMENTARY MATERIALS



Supplementary Figure 1: Identification of differentially expressed genes for HNSC, LIHC, READ and UCEC. Venn diagram showing the overlap between differentially expressed genes selected by each method.



Supplementary Figure 2: Cell type enrichment of co-expression modules. The cell-type enrichment analysis of cluster 2 (upper left-hand corner), cluster 5 (upper right-hand corner), cluster 7 (bottom) is shown in as -log10 (Benjamini-Hochberg corrected p-value).



Supplementary Figure 3: Survival analysis of the cluster 2 for 8 types of cancers. Kaplan-Meier curves are used in survival analysis. The calculation of the log-rank p-values are based on the split of patient groups using non-negative matrix factorization. The number of patients in each group is also labeled in each panel.



Supplementary Figure 4: Survival analysis of the cluster 7 for 8 types of cancers. Kaplan-Meier curves are used in survival analysis. The calculation of the log-rank p-values are based on the split of patient groups using non-negative matrix factorization. The number of patients in each group is also labeled in each panel.



Supplementary Figure 5: Survival analysis of the cluster 8 for 8 types of cancers. Kaplan-Meier curves are used in survival analysis. The calculation of the log-rank p-values are based on the split of patient groups using non-negative matrix factorization. The number of patients in each group is also labeled in each panel.



Supplementary Figure 6: Survival analysis of the cluster 5 for other 8 types of cancers. Kaplan-Meier curves are used in survival analysis for GBM, LUSC, OV, CESC, KIRP, ACC, SKCM and LGG. The calculation of the log-rank p-values are based on the split of patient groups using non-negative matrix factorization. The number of patients in each group is also labeled in each panel.

| Module | Genes in the cluster | Genes in the FI network | | |
|------------|----------------------|-------------------------|--|--|
| Cluster 2 | 160 | 129 | | |
| Cluster 5 | 74 | 62 | | |
| Cluster 7 | 94 | 72 | | |
| Cluster 8 | 375 | 327 | | |
| Cluster 10 | 30 | | | |

Supplementary Table 1: Number of genes in the cluster as well as number of genes used in the corresponding FI network

Four clusters were identified in the eight cancer types.

Supplementary Table 2: GO BPs of the four clusters

See Supplementary File 1

Supplementary Table 3: Number of genes in the cluster

| Module | Genes in the cluster | | |
|-----------|----------------------|--|--|
| Cluster 4 | 279 | | |
| Cluster 6 | 128 | | |

Two clusters were identified in the four cancer types.

Supplementary Table 4: Cell type enrichment of the four clusters. The score is-log10 (Benjamini-Hochberg corrected p-value)

See Supplementary File 2

Supplementary Table 5: The enriched diseases in the four clusters

| | Disease | Benjamini-Hochberg corrected p-value |
|-----------|--------------------------------|--------------------------------------|
| Cluster 2 | Neovascularization, Pathologic | 7.23E-23 |
| | Neoplastic Processes | 5.97E-22 |
| | Carcinoma | 5.97E-22 |
| | Neoplasms | 1.32E-19 |
| | Metaplasia | 3.94E-19 |
| | Adhesion | 1.93E-17 |
| | epithelial cancers | 4.32E-17 |
| | Neoplasm Metastasis | 4.32E-17 |
| | Vascular Diseases | 4.16E-16 |
| | cancer or viral infections | 1.36E-15 |
| Cluster 5 | cancer or viral infections | 1.17E-10 |
| | Neoplasms | 1.55E-10 |
| | Carcinoma | 5.07E-10 |
| | epithelial cancers | 5.26E-10 |
| | Urogenital Neoplasms | 1.18E-08 |
| | Collagen Diseases | 1.59E-08 |
| | Chorioamnionitis | 2.04E-08 |
| | Prostatic Neoplasms | 2.65E-08 |
| | Neoplastic Processes | 7.08E-08 |
| | Male Urogenital Diseases | 8.39E-08 |
| Cluster 7 | Inflammation | 1.68E-15 |
| | Immune System Diseases | 7.12E-15 |
| | Chronic Disease | 2.56E-11 |
| | Common Cold | 2.62E-11 |
| | Respiratory Tract Infections | 2.62E-11 |
| | Autoimmune Diseases | 3.01E-11 |
| | Arthritis, Rheumatoid | 4.31E-11 |
| | Hypersensitivity | 3.03E-10 |
| | Infection | 3.83E-10 |
| | Joint Diseases | 8.06E-10 |
| Cluster 8 | Neoplastic Processes | 4.48E-36 |
| | Neoplasm Metastasis | 1.99E-31 |
| | cancer or viral infections | 5.98E-30 |
| | Neoplasm Invasiveness | 7.15E-29 |
| | Carcinoma | 1.44E-28 |
| | Neoplasms | 6.39E-28 |
| | Fibrosis | 4.27E-27 |
| | Adhesion | 8.04E-25 |
| | Collagen Diseases | 1.02E-24 |
| | Adenocarcinoma | 9.35E-23 |

Supplementary Table 6: The enriched miRNAs in the four clusters

See Supplementary File 3

Supplementary Table 7: The enriched transcription factor targets (TFT) in the four clusters

See Supplementary File 4

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| M1 | M2 | M3 | M4 | M5 | M6 | M7 | M8 | M9 |
|----------|---------|---------|---------|------------|-------|------|-------|---------|
| ACSM3 | ADAM12 | ALDH1A3 | ACTG2 | GALNT6 | AMACR | DPP4 | EFHD1 | MXRA5 |
| ANO7 | CHAD | CA12 | ARHGAP6 | ST6GALNAC2 | FOLH1 | FAP | | S100A16 |
| CRABP2 | COL11A1 | СКВ | CNN1 | | | VCAN | | FNDC1 |
| CTHRC1 | COL1A1 | GDF15 | GFRA1 | | | | | TMPRSS2 |
| FABP4 | COL1A2 | MAOA | MMP11 | | | | | PGM5 |
| FAM3B | COL3A1 | PRUNE2 | MYH11 | | | | | PDE9A |
| GATA3 | COL5A2 | S100A14 | SFRP2 | | | | | TBC1D9 |
| GPRC5A | COL6A3 | SLC2A12 | | | | | | HGD |
| KLF15 | DES | SPON2 | | | | | | MIR614 |
| KRT19 | ECM1 | STC2 | | | | | | LRRC15 |
| KRT7 | FN1 | STEAP1 | | | | | | STEAP2 |
| MFAP2 | PLAU | TRPS1 | | | | | | PCP4 |
| MFAP5 | POSTN | | | | | | | |
| NEFH | THBS2 | | | | | | | |
| NKX3-1 | | | | | | | | |
| PLA1A | | | | | | | | |
| SERPINA5 | | | | | | | | |
| SLC4A4 | | | | | | | | |
| TFAP2A | | | | | | | | |
| TSPAN1 | | | | | | | | |
| WFDC1 | _ | | | | | | | |

Supplementary Table 8: The genes in the nine sub-modules

| Abbreviated cancer name | Tumor samples | Normal samples |
|-------------------------|----------------------|----------------|
| BRCA | 1119 | 113 |
| KIRC | 542 | 72 |
| PRAD | 502 | 52 |
| THCA | 513 | 59 |
| LUAD | 541 | 59 |
| BLCA | 414 | 19 |
| COAD | 483 | 41 |
| STAD | 420 | 37 |
| UCEC | 554 | 35 |
| HNSC | 504 | 44 |
| READ | 167 | 10 |
| LIHC | 374 | 50 |

Supplementary Table 9: The used cancer types and number of the corresponding samples