

Figure S1 OS results of immune cells are significantly different between groups (determined by the median value). OS, overall survival.

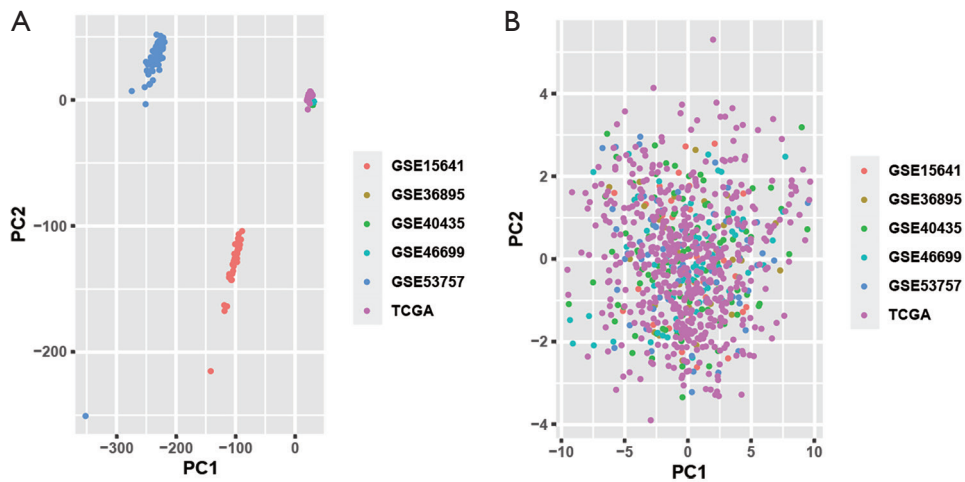


Figure S2 Two-dimensional plots are shown of principal components calculated PCA. (A) PCA of the expression matrix of six different datasets. (B) PCA of the immune cells enrichment scores of six different datasets. PCA, principal components analysis; TCGA, The Cancer Genome Atlas.

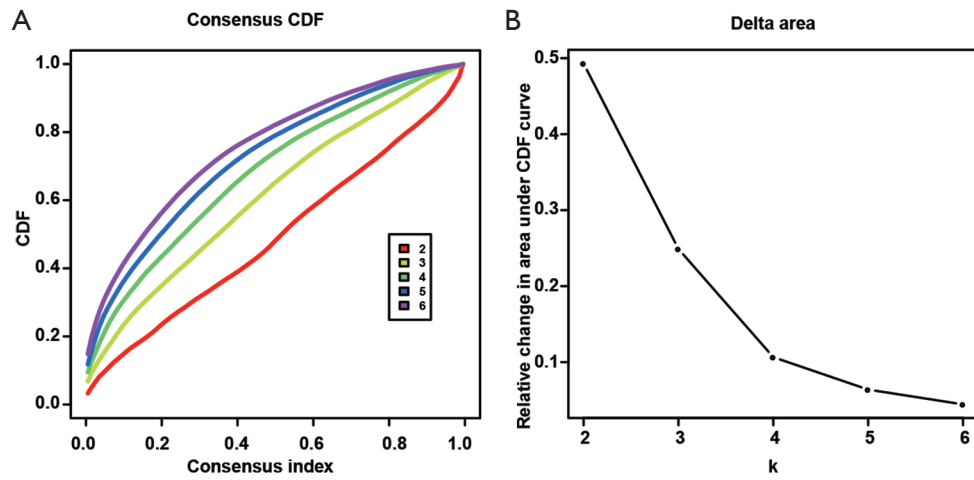


Figure S3 The selection of immune subtype numbers. (A) CC CDF for $k=2$ to 6. (B) Delta area curve of CC, indicating the relative change in area under CDF curve for each category number k compared with $k-1$. The horizontal axis represents the category number k , and the vertical axis represents the relative change in area under the CDF curve. CC, consensus clustering; CDF, cumulative distribution function.

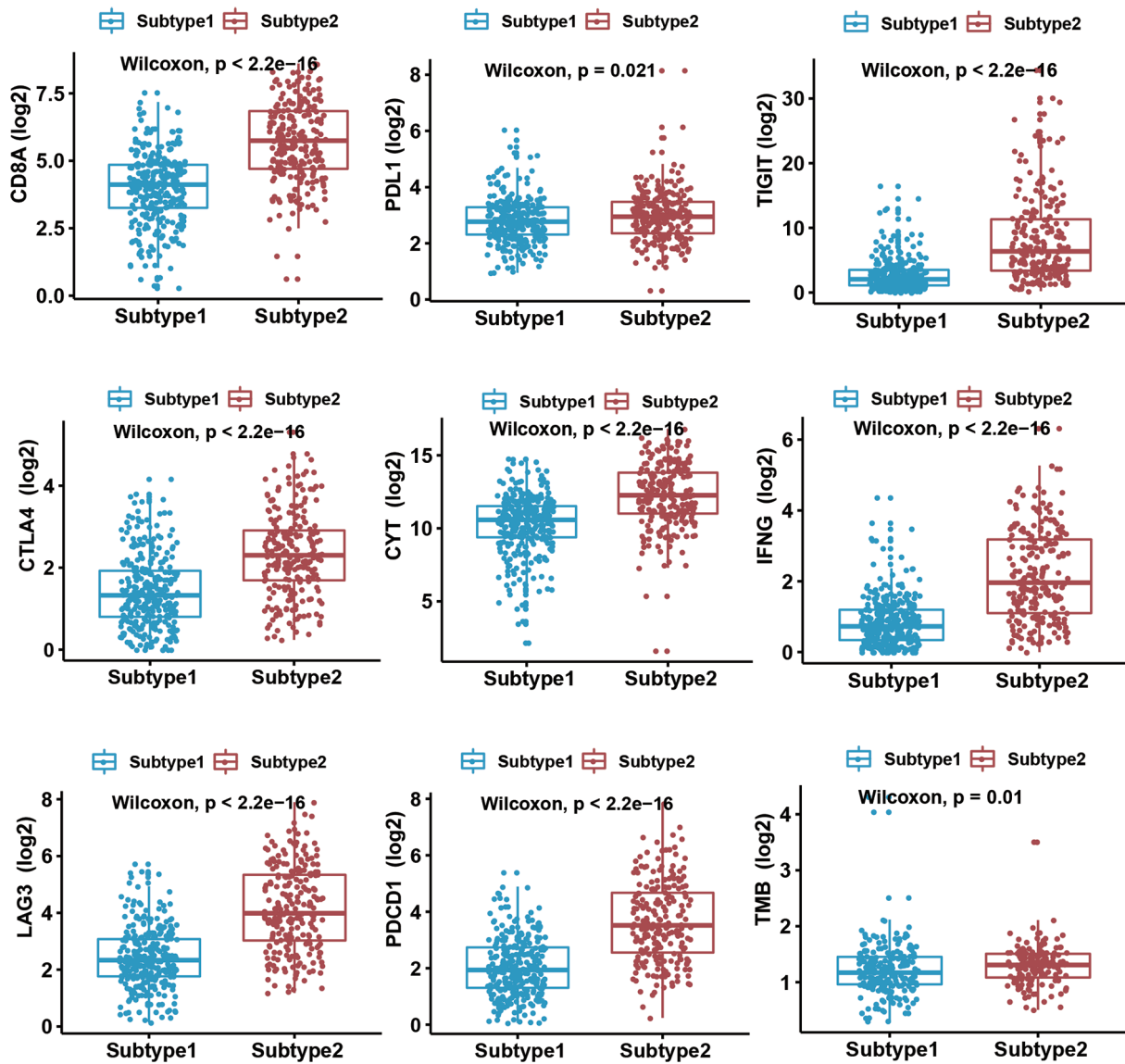


Figure S4 Comparison of the tumor immunotherapy indicators between the two immune subtypes in the TCGA dataset. Subtype2 tumors had significantly higher *CD8A*, *PDL1*, *TIGIT*, *CTLA4*, *CYT*, *IFNG*, *LAG3*, *PD1* (*PDCD1*) and *TMB* than subtype1 tumors ($P < 0.05$). TCGA, The Cancer Genome Atlas; TMB, tumor mutational burden.

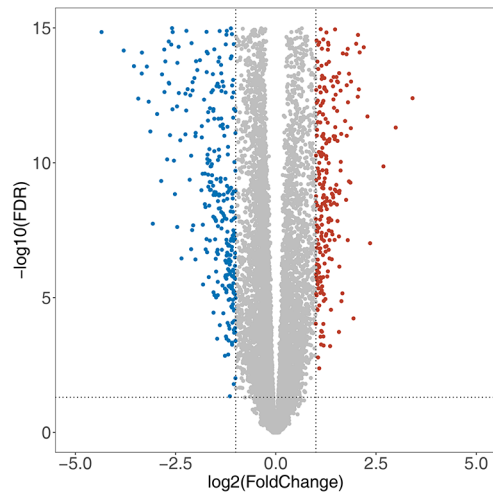


Figure S5 Volcano plot showing the gene expression differences between immune subtypes. Blue dots, down-regulated genes in subtype2. Red dots, upregulated genes in subtype2. FDR, false discovery rate.

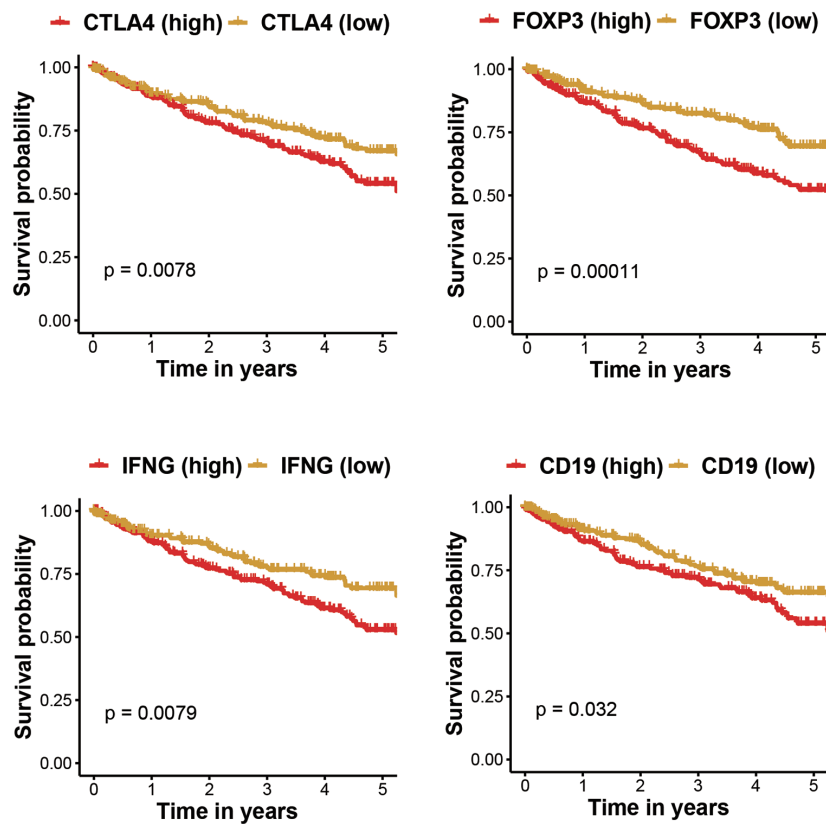


Figure S6 OS results of four hub genes in the turquoise module are significantly different between groups (determined by the median value). OS, overall survival.

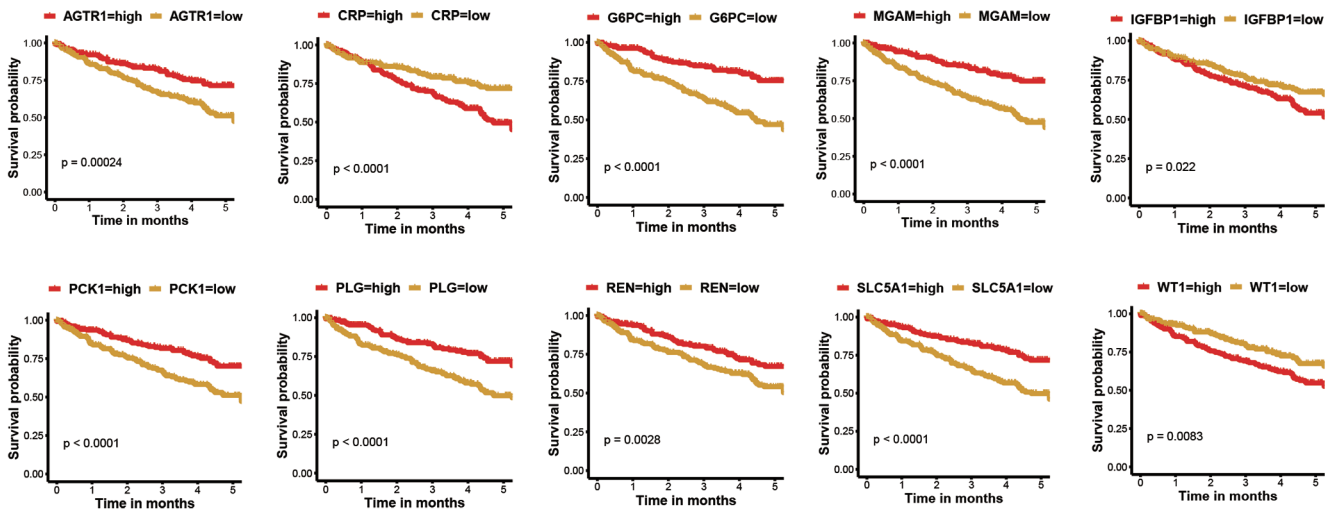


Figure S7 OS results of ten hub genes in the blue module are significantly different between groups (determined by the median value). OS, overall survival.

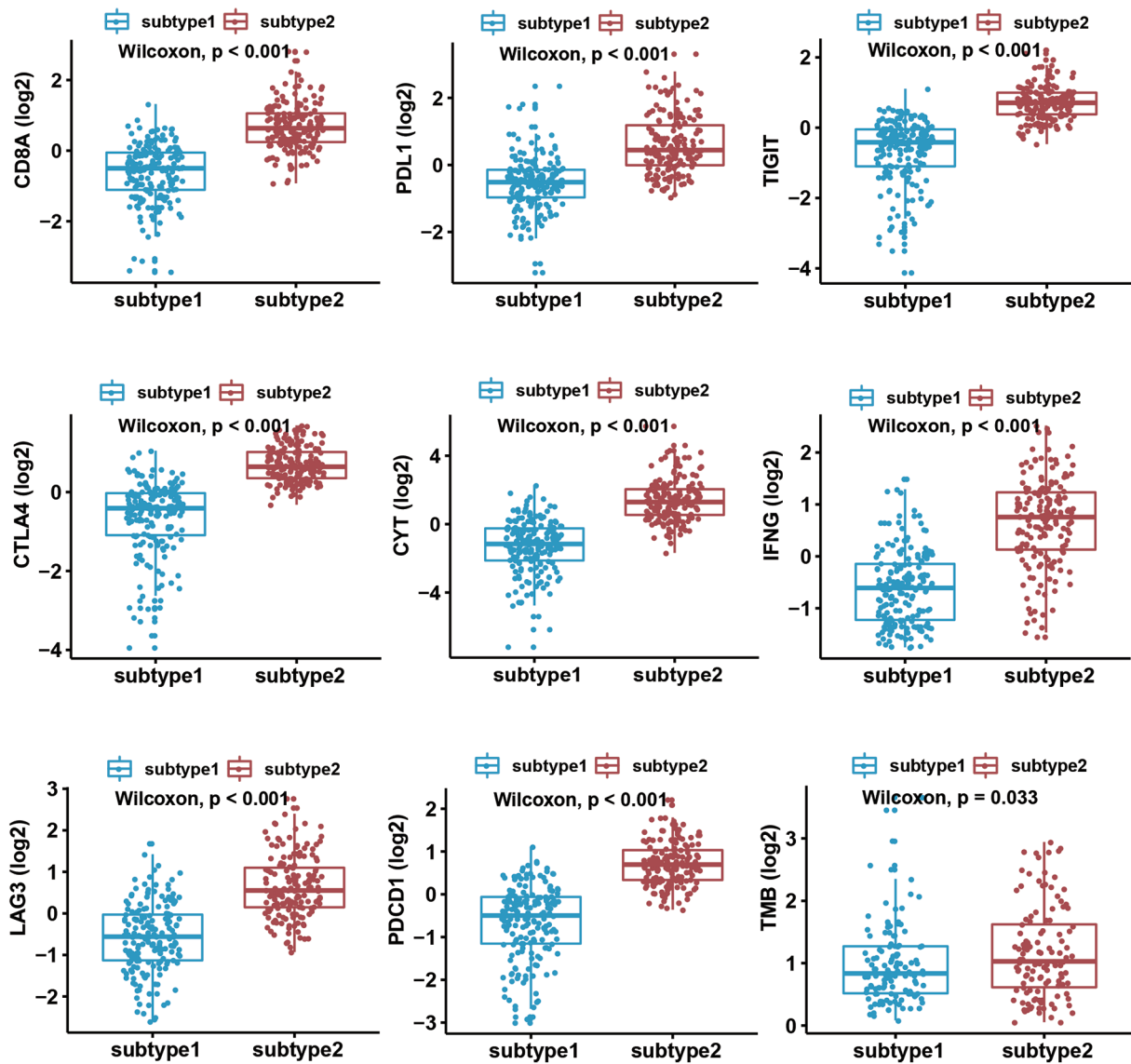


Figure S8 Comparison of the tumor immunotherapy indicators between the two immune subtypes in the IMvigor210 dataset. Subtype2 tumors had significantly higher *CD8A*, *PDL1*, *TIGIT*, *CTLA4*, *CYT*, *IFNG*, *LAG3*, *PD1* (*PDCD1*) and *TMB* than subtype1 tumors ($P < 0.05$). TMB, tumor mutational burden.

Table S1 The enriched pathways in immune subtype1

| Pathway | Padj | NES | Size |
|---|-------|--------------|------|
| PPAR signaling pathway | <0.05 | -2.076462405 | 69 |
| Oxidative phosphorylation | <0.05 | -2.075496402 | 116 |
| Vibrio cholerae infection | <0.05 | -2.01780083 | 52 |
| Fatty acid metabolism | <0.05 | -1.99482005 | 42 |
| Retinol metabolism | <0.05 | -1.927153616 | 63 |
| Tyrosine metabolism | <0.05 | -1.908248892 | 41 |
| Drug metabolism cytochrome P450 | <0.05 | -1.829246779 | 70 |
| Glycolysis gluconeogenesis | <0.05 | -1.793530881 | 61 |
| Propanoate metabolism | <0.05 | -1.779770488 | 32 |
| Epithelial cell signaling in <i>Helicobacter pylori</i> infection | <0.05 | -1.773782491 | 67 |

A negative NES means that genes over-represented in the gene set are upregulated in immune subtype1. Padj, adjusted P values (the FDR); FDR, false discovery rate; NES, normalized enrichment score.

Table S2 The enriched pathways in immune subtype2

| Pathway | Padj | NES | Size |
|--|-------|-------------|------|
| Natural killer cell mediated cytotoxicity | <0.05 | 2.101136753 | 130 |
| Leishmania infections | <0.05 | 2.112326976 | 69 |
| Asthma | <0.05 | 2.147862606 | 28 |
| T cell receptor signaling pathway | <0.05 | 2.203600853 | 106 |
| Chemokine signaling pathway | <0.05 | 2.213683506 | 184 |
| Allograft rejection | <0.05 | 2.244700522 | 35 |
| Graft versus host disease | <0.05 | 2.25556481 | 37 |
| Primary immunodeficiency | <0.05 | 2.271887799 | 35 |
| Cytokine-cytokine receptor interaction | <0.05 | 2.28028487 | 257 |
| Intestinal immune network for IgA production | <0.05 | 2.310500831 | 46 |
| Antigen processing and presentation | <0.05 | 2.363167045 | 79 |

A positive NES means that genes over-represented in the gene set are upregulated in immune subtype2. Padj, adjusted P values (the FDR); FDR, false discovery rate; NES, normalized enrichment score.

Table S3 Baseline characteristics of patients in the IMvigor210 cohort

| Clinical type | Sample [%] |
|-----------------------|------------|
| Immunotherapy outcome | |
| CR/PR | 68 [19] |
| SD/PD | 230 [66] |
| NA | 50 [15] |
| Gender | |
| Male | 272 [78] |
| Female | 76 [22] |
| Tobacco history | |
| Previous | 197 [57] |
| Never | 116 [33] |
| Current | 35 [10] |
| Received platinum | |
| Yes | 272 [78] |
| No | 76 [22] |

CR, complete response; PR, partial response; SD, stable disease; PD, progressive disease; NA, data is not available.