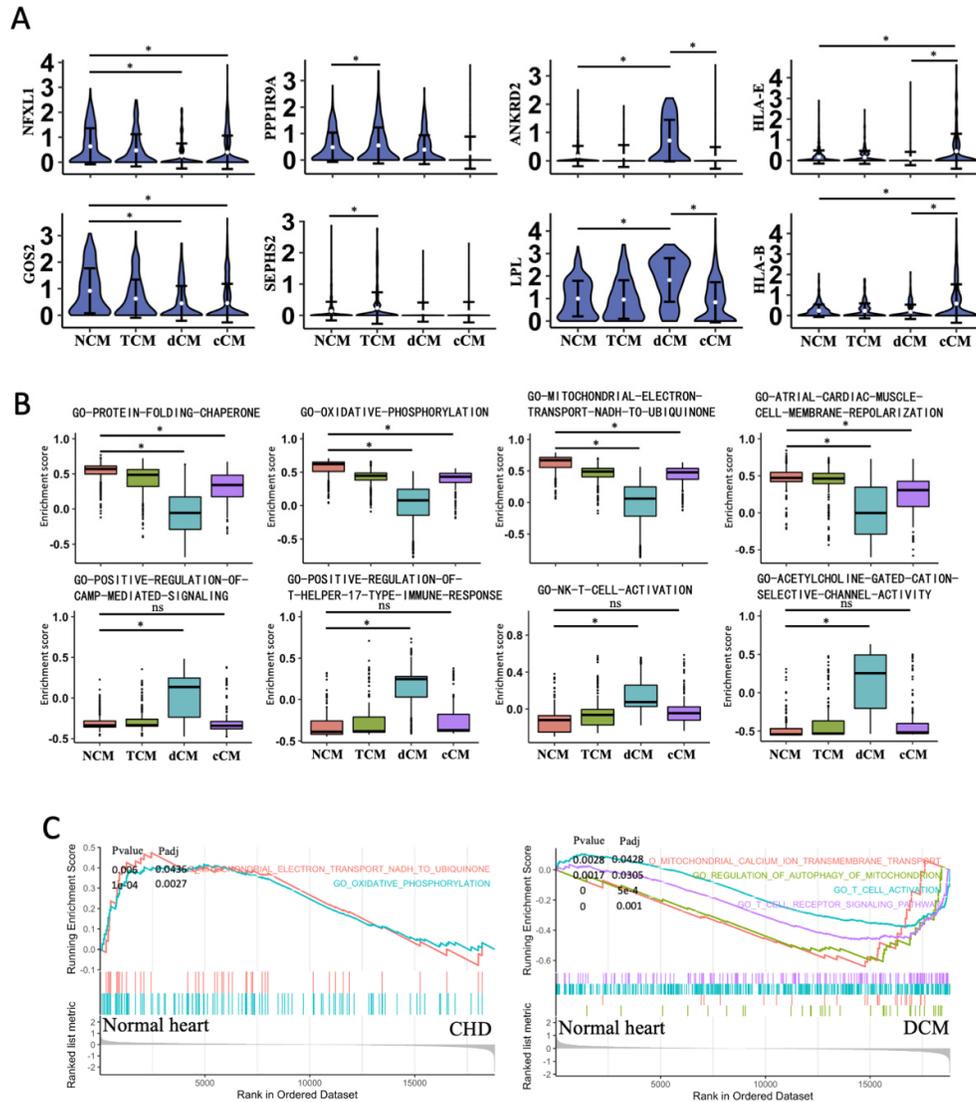
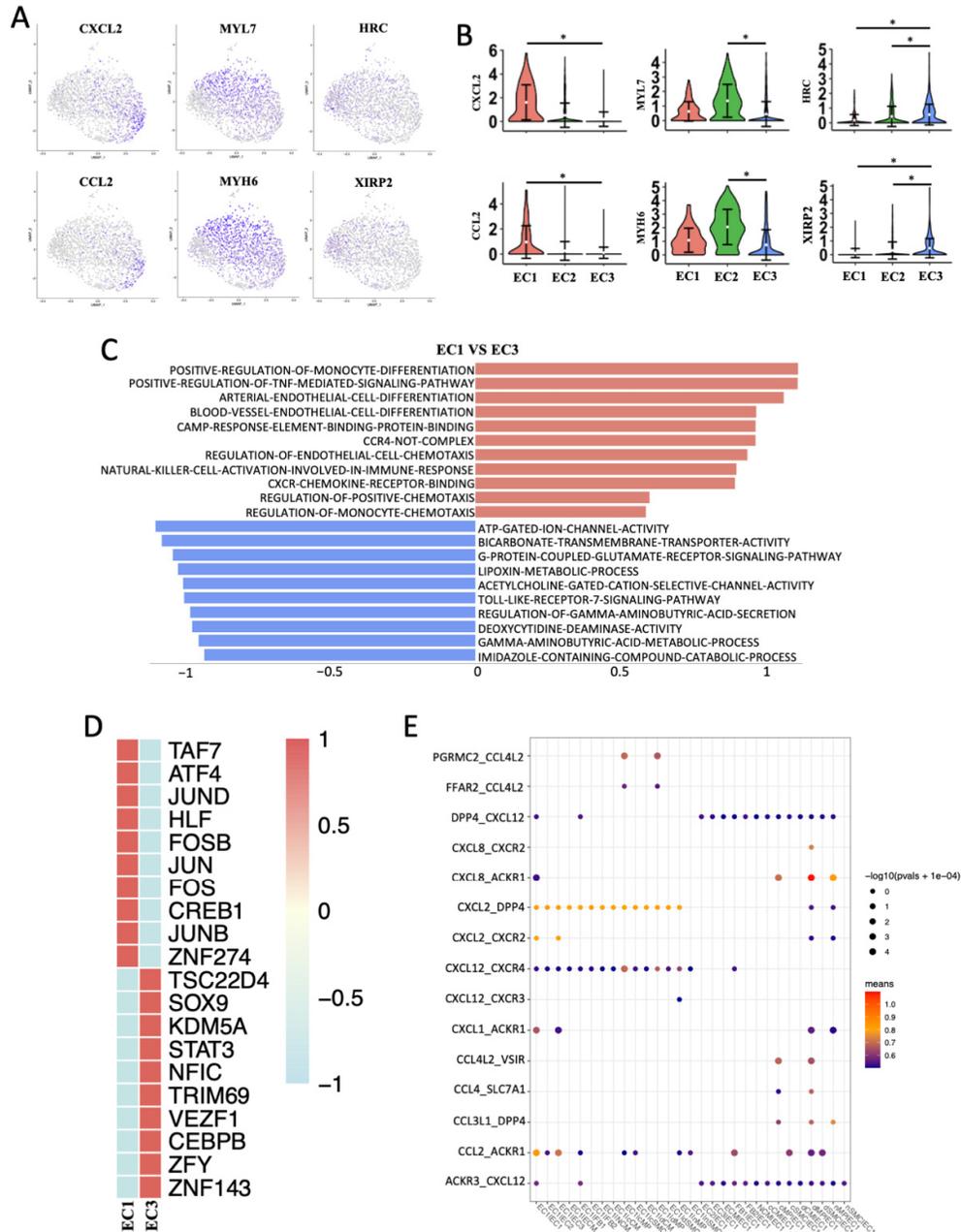


**Supplementary Figure S1. Five cell types were identified from Single-cell transcriptome of heart disease. (A)** UMAP plots of five cell types, clusters are color-coded by cell sample. **(B)** proportion of five cell types in each sample. **(C)** UMAP plot of each sample (cHF, dHF and nHF), colored by cell types.

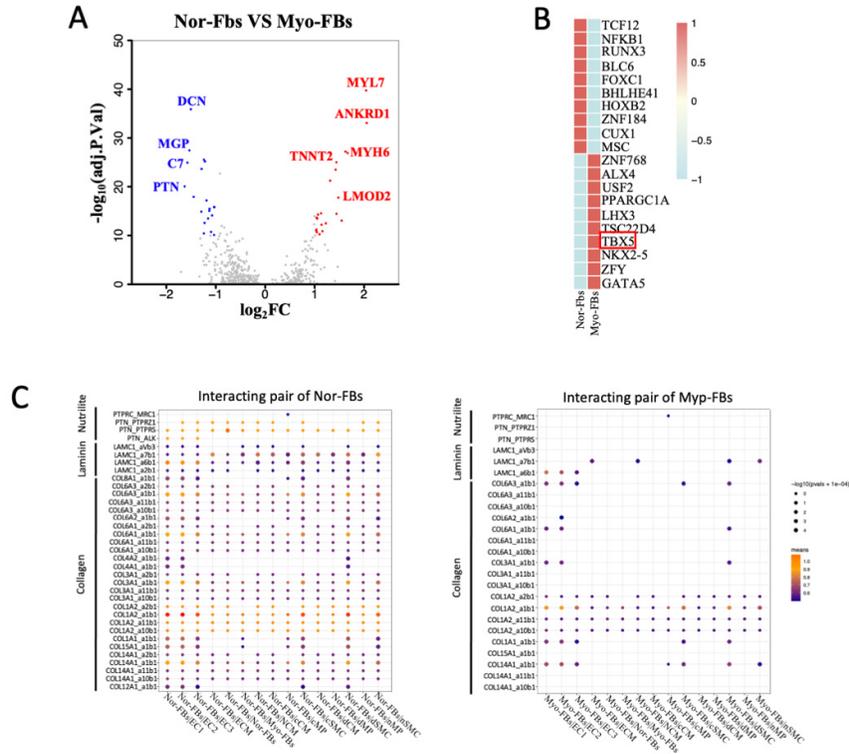


**Supplementary Figure S2. Cardiomyocytes (CMs) cluster into 4 distinct phenotypes.** (A) violin plots indicate differentially expressed genes in phenotypes of CMs. \* $P_{adj} < 0.05$  Wilcoxon test. (B) Multiple pathway enrichments show the characteristics of the cCM and dCM, as well as the difference between cCM and dCM. Boxplot y-axis demonstrates enrichment score of pathways. \* $p_{adj} < 0.05$  Wilcoxon test. (C) GSEA results showed pathway difference between normal heart and ischemic or dilated cardiomyopathy.



**Supplementary Figure S3. Endothelial cells in HF have two special disease-related phenotypes.** (A) Expression of marker genes for the phenotypes of ECs defined above each panel. (B) The expression of known cell type discriminating genes in EC1, EC2 and EC3 from 3 samples. \* $P_{adj} < 0.05$  Wilcox test. (C) Differences in pathway

enrichment scored by GSVA between EC1 and EC3. **(D)** Differences in TFs active scored by SCENIC between EC1 and EC3. **(E)** Immunity-related interacting pair between EC1 and other cell clusters.



**Supplementary Figure S4. Fibroblasts transformed into Myofibroblasts. (A)** Volcano show the DEGs ( $\log_2FC > 1$ ) between Nor-FBs and Myo-FBs. **(B)** Differences in TFs active scored by SCENIC between Nor-FBs and Myo-FBs. **(C)** Crosstalk of Nor-FBs or Myo-FBs showed the functional change during the phenotypic transformation.