

SUPPLEMENTAL DATA

**Deciphering transcriptional dynamics of
cardiac hypertrophy and failure in a
chamber-specific manner**

Sample ID	Tissues	raw_reads number	clean_reads number	clean_bases	error_rate	Q30	GC (%)
A_CHF1	atrium	51086866	50601886	7.59G	0.03	92.87	48.83
A_CHF2	atrium	46548484	46108434	6.92G	0.03	93.04	48.57
A_CHF3	atrium	47358694	46925514	7.04G	0.03	93.12	48.89
A_HCM1	atrium	46677678	46296298	6.94G	0.03	93.14	48.91
A_HCM2	atrium	46908252	46563004	6.98G	0.03	92.55	49.07
A_HCM3	atrium	49608912	49187858	7.38G	0.03	93.37	48.24
A_sham1	atrium	45706142	45025986	6.75G	0.03	93.34	48.79
A_sham2	atrium	45489412	44822164	6.72G	0.03	93.37	48.73
A_sham3	atrium	47544878	47048378	7.06G	0.03	93.14	48.91
LV_CHF1	left ventricle	46716808	46195920	6.93G	0.03	92.58	47.18
LV_CHF2	left ventricle	46182796	45628840	6.84G	0.03	93.28	47.3
LV_CHF3	left ventricle	46450658	45837024	6.88G	0.03	92.95	47.16
LV_HCM1	left ventricle	46476284	45909980	6.89G	0.03	92.8	47.62
LV_HCM2	left ventricle	47613930	47157086	7.07G	0.03	93.09	47.41
LV_HCM3	left ventricle	45201762	44682114	6.7G	0.03	93.18	47.34
LV_sham1	left ventricle	46297952	45844554	6.88G	0.03	92.76	47.03
LV_sham2	left ventricle	47473316	46840548	7.03G	0.03	92.98	44.41
LV_sham3	left ventricle	46983382	46527468	6.98G	0.03	93.03	47.2
RV_CHF1	right ventricle	47248666	46444388	6.97G	0.03	92.16	40.51
RV_CHF2	right ventricle	45278016	43648060	6.55G	0.03	92.42	44.66
RV_CHF3	right ventricle	47637072	46997014	7.05G	0.03	92.37	41.72
RV_HCM1	right ventricle	47051652	46289232	6.94G	0.03	92.61	40.18
RV_HCM2	right ventricle	47109924	46395186	6.96G	0.03	92.23	40.17
RV_HCM3	right ventricle	46032126	45054048	6.76G	0.03	91.71	40.51
RV_sham1	right ventricle	51678400	50798664	7.62G	0.03	91.48	41.95
RV_sham2	right ventricle	47078410	46588702	6.99G	0.03	92.62	41.51
RV_sham3	right ventricle	46801706	45751680	6.86G	0.03	91.79	40.73

Table S1 Sample information and sequencing statistics

LA: left atrium; LV: left ventricle; RV: right ventricle

	CH	HF
	muscle contraction extracellular matrix organization extracellular structure organization external encapsulating structure organization muscle system process collagen-containing extracellular matrix micturition transmembrane receptor protein STK signaling pathway	circadian rhythm; rhythmic process hydrogen peroxide metabolic/catabolic process circadian behavior; rhythmic behavior; locomotor rhythm circadian regulation of gene expression oxygen transport; gas transport haptoglobin-hemoglobin complex distal axon; myelin sheath; terminal bouton presynapse; neurofilament; axon terminus neuron projection terminus extrinsic component of membrane intermediate filament oxygen carrier activity; oxygen binding organic acid binding; phospholipase binding cell adhesion molecule binding; nuclear receptor binding calcium-activated potassium channel activity peroxidase activity; oxidoreductase activity structural constituent of cytoskeleton
LA	excretion regulation of smooth muscle contraction phasic smooth muscle contraction neuronal cell body membrane cell body membrane dendrite membrane dendritic spine membrane	
LV	muscle contraction extracellular matrix organization extracellular structure organization	fat cell differentiation extracellular matrix organization extracellular structure organization

<p>external encapsulating structure organization muscle system process collagen-containing extracellular matrix regulation of cardiac muscle cell membrane repolarization cardiac muscle cell membrane repolarization regulation of membrane repolarization membrane repolarization regulation of heart rate chromosome, centromeric region condensed chromosome voltage-gated sodium channel complex myosin filament chromosomal region sodium channel complex</p>	<p>external encapsulating structure organization extracellular matrix structural constituent collagen-containing extracellular matrix amino acid transmembrane transport regulation of amino acid import across plasma membrane regulation of amino acid transmembrane transport regulation of cardiac muscle cell apoptotic process segmentation; collagen trimer regulation of striated muscle cell apoptotic process C-C chemokine receptor activity; C-C chemokine binding G protein-coupled chemoattractant receptor activity chemokine receptor activity; cytokine receptor activity cytokine binding; chemokine binding proton-transporting ATP synthase activity, rotational mechanism immune receptor activity</p>
<p>negative regulation of transmembrane receptor protein STK signaling pathway extracellular matrix organization extracellular structure organization RV external encapsulating structure organization regulation of transmembrane receptor protein STK signaling pathway respiratory electron transport chain L-glutamate transmembrane transporter activity</p>	<p>striated muscle cell differentiation extracellular matrix organization extracellular structure organization external encapsulating structure organization collagen-containing extracellular matrix extracellular matrix structural constituent muscle cell differentiation; muscle cell development</p>

retinoic acid 4-hydroxylase activity	regulation of striated muscle cell differentiation
C4-dicarboxylate transmembrane transporter activity	myotube differentiation
retinoic acid binding	entrainment of circadian clock by photoperiod
NAD(P)H oxidoreductase activity	sarcomere; myofibril; contractile fiber
acidic amino acid transmembrane transporter activity	I band; Z disc
heme binding	heparin binding
tetrapyrrole binding	sulfur compound binding; fibronectin binding
heparin binding	photoperiodism; glycosaminoglycan binding

Table S2 Top 10 BP, CC, and MF GO terms in CH and HF, respectively

Note: GO terms with a red background denote Gene Ontology (GO) terms enriched in both the LA and LV, GO terms with a blue background represent GO terms shared between the LA and RV, and yellow in LV and RV. Without any background color, unique GO terms are enriched in each chamber; for example, no shared GO terms between the atrium and left ventricle.

Entries in bold denote GO terms enriched in CH and HF in the LA, LV, and RV, respectively.

CH- cardiac hypertrophy, HF- heart failure, LA- left atrium, LV- left ventricle, RV- right ventricle, BP- biological process, CC- cellular component, MF- molecular function;

	CH	HF
	Cytokine-cytokine receptor interaction	Cytokine-cytokine receptor interaction
	Vascular smooth muscle contraction	Vascular Smooth Muscle Contraction
	Salivary secretion	Circadian Rhythm
	Calcium signaling pathway	African trypanosomiasis
	Pyruvate metabolism	NF-kappa B signaling pathway
	Neuroactive ligand-receptor interaction	Malaria
LA	Drug metabolism - cytochrome P450	Hippo signaling pathway
	Ras signaling pathway	
	Metabolism of xenobiotics by cytochrome P450	
	Butanoate metabolism	
	Fatty acid degradation	
	Tryptophan metabolism	
	Glutamatergic synapse	
	cAMP signaling pathway	
	Cytokine-cytokine receptor interaction	Cytokine-cytokine receptor interaction
	Chemokine signaling pathway	Chemokine signaling pathway
	Adrenergic signaling in cardiomyocytes	Adrenergic signaling in cardiomyocytes
	Aldosterone synthesis and secretion	Aldosterone synthesis and secretion
	Protein digestion and absorption	Platelet activation
LV	Platelet activation	Cardiac muscle contraction
	Cardiac muscle contraction	Protein digestion and absorption
	Cell cycle	Vascular Smooth Muscle Contraction
	Progesterone-mediated oocyte maturation	Circadian rhythm
	Oocyte meiosis	Oxytocin signaling pathway
	ECM-receptor interaction	Viral protein interaction with cytokine and cytokine receptor

Diabetic cardiomyopathy	Circadian entrainment
Homologous recombination	cGMP-PKG signaling pathway
p53 signaling pathway	Dilated cardiomyopathy
	Glycosphingolipid biosynthesis - ganglio series
Circadian entrainment	Circadian entrainment
Oxytocin signaling pathway	Oxytocin signaling pathway
African trypanosomiasis	Vascular Smooth Muscle Contraction
Adrenergic signaling in cardiomyocytes	Circadian Rhythm
Natural killer cell mediated cytotoxicity	Protein digestion and absorption
Mucin type O-glycan biosynthesis	cGMP-PKG signaling pathway
Primary immunodeficiency	Viral protein interaction with cytokine and cytokine receptor
RV MAPK signaling pathway	ECM-receptor interaction
T cell receptor signaling pathway	Estrogen signaling pathway
Intestinal immune network for IgA production	Insulin secretion
One carbon pool by folate	Human papillomavirus infection
Cholesterol metabolism	TNF signaling pathway
Vitamin digestion and absorption	Ovarian steroidogenesis
Malaria	Calcium signaling pathway
	Drug metabolism - other enzymes

Table S3 The top 15 KEGG pathways in CH and HF

Note: KEGG pathway terms with a red background denote the KEGG pathway enriched in both the LA and LV, KEGG pathways with a blue background represent the KEGG pathway shared between the LA and RV, and yellow in LV and RV. Without any background color, unique KEGG pathways are enriched in each chamber.

KEGG pathways in bold denote the KEGG pathway enriched in CH and HF in the LA, LV, and RV, respectively.

CH- cardiac hypertrophy, HF- heart failure, LA- left atrium, LV- left ventricle, RV- right ventricle, BP- biological process, CC- cellular component, MF- molecular function, KEGG- Kyoto Encyclopedia of Genes and Genomes;

	CH		HF
LA	Degradation of the extracellular matrix Extracellular matrix organization Assembly of collagen fibrils and other multimeric structures Collagen formation Collagen biosynthesis and modifying enzymes Crosslinking of collagen fibrils Integrin cell surface interactions Non-integrin membrane-ECM interactions Collagen degradation Collagen chain trimerization Muscle contraction Striated Muscle Contraction ECM proteoglycans NCAM1 interactions NCAM signaling for neurite out-growth		Erythrocytes take up oxygen and release carbon dioxide Erythrocytes take up carbon dioxide and release oxygen O2/CO2 exchange in erythrocytes Acetylcholine Neurotransmitter Release Cycle Scavenging of heme from plasma Adherens junctions interactions Neuronal System Potassium Channels Phase I - Functionalization of compounds Binding and Uptake of Ligands by Scavenger Receptors Muscle contraction Striated Muscle Contraction O-glycosylation of TSR domain-containing proteins Xenobiotics Attenuation phase
LV	Extracellular matrix organization Assembly of collagen fibrils and other multimeric structures Collagen formation Integrin cell surface interactions Degradation of the extracellular matrix		Assembly of collagen fibrils and other multimeric structures Extracellular matrix organization Collagen formation Collagen biosynthesis and modifying enzymes Integrin cell surface interactions

<p>Collagen biosynthesis and modifying enzymes Collagen degradation Collagen chain trimerization Crosslinking of collagen fibrils Non-integrin membrane-ECM interactions</p> <p>Laminin interactions Elastic fibre formation Signaling by Rho GTPases Kinesins Anchoring fibril formation</p>	<p>Crosslinking of collagen fibrils Collagen degradation Collagen chain trimerization Non-integrin membrane-ECM interactions Degradation of the extracellular matrix Laminin interactions Axon guidance Developmental Biology NCAM signaling for neurite out-growth NCAM1 interactions</p>
<p>RV</p> <p>Respiratory electron transport Complex I biogenesis The citric acid (TCA) cycle and respiratory electron transport Respiratory electron transport, ATP synthesis by chemiosmotic coupling, and heat production by uncoupling proteins. Phase 0 - rapid depolarisation Cardiac conduction</p> <p>Erythrocytes take up carbon dioxide and release oxygen O₂/CO₂ exchange in erythrocytes Sema4D in semaphorin signaling Signaling by Retinoic Acid</p>	<p>Extracellular matrix organization Integrin cell surface interactions Assembly of collagen fibrils and other multimeric structures Collagen formation Collagen biosynthesis and modifying enzymes</p> <p>Nonsense Mediated Decay (NMD) enhanced by the Exon Junction Complex (EJC) Nonsense-Mediated Decay (NMD) O-linked glycosylation ECM proteoglycans Anchoring fibril formation</p>

GPVI-mediated activation cascade	MET activates PTK2 signaling
Collagen degradation	MET promotes cell motility
Collagen chain trimerization	Platelet degranulation
Muscle contraction	O-glycosylation of TSR domain-containing proteins
Striated Muscle Contraction	Striated Muscle Contraction

Table S4 The top 15 Reactome pathways in CH and HF

Note: Reactome pathway terms with a red background denote the Reactome pathway enriched in both the LA and LV, Reactome pathways with a blue background represent the Reactome pathway shared between the LA and RV, and yellow in LV and RV. Without any background color, unique Reactome pathways are enriched in each chamber.

Reactome pathways in bold denote the Reactome pathway enriched in CH and HF in the LA, LV, and RV, respectively.

CH- cardiac hypertrophy, HF- heart failure, LA- left atrium, LV- left ventricle, RV- right ventricle, BP- biological process, CC- cellular component, MF- molecular function;

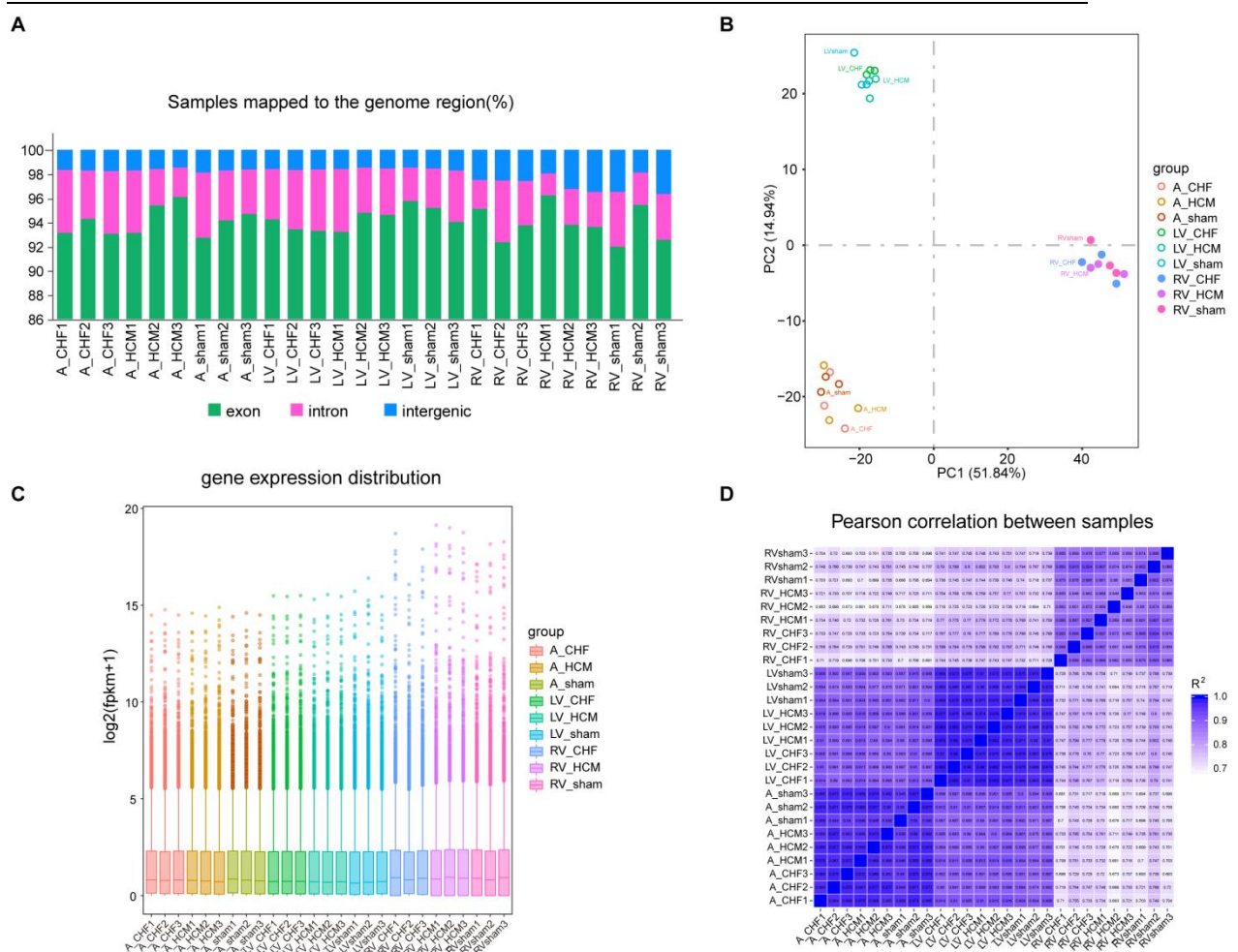


Figure S1 Quality control and quantitative analysis of sequencing data from all samples. (A) Reads of all samples were aligned to the reference genome. (B) PCA using FPKM of samples. The samples were clustered according to the sampling location. (C) Boxplots of gene expression distribution. The X-axis is the name of the samples; Y-axis is log₂ (FPKM+1). (D) Correlation of gene expression levels among samples based on Pearson correlation between samples. Atrium (left), LV- left ventricle, RV- right ventricle, PCA- principal component analysis, FPKM- Fragments Per Kilobase of transcript sequence per Millions base pairs;

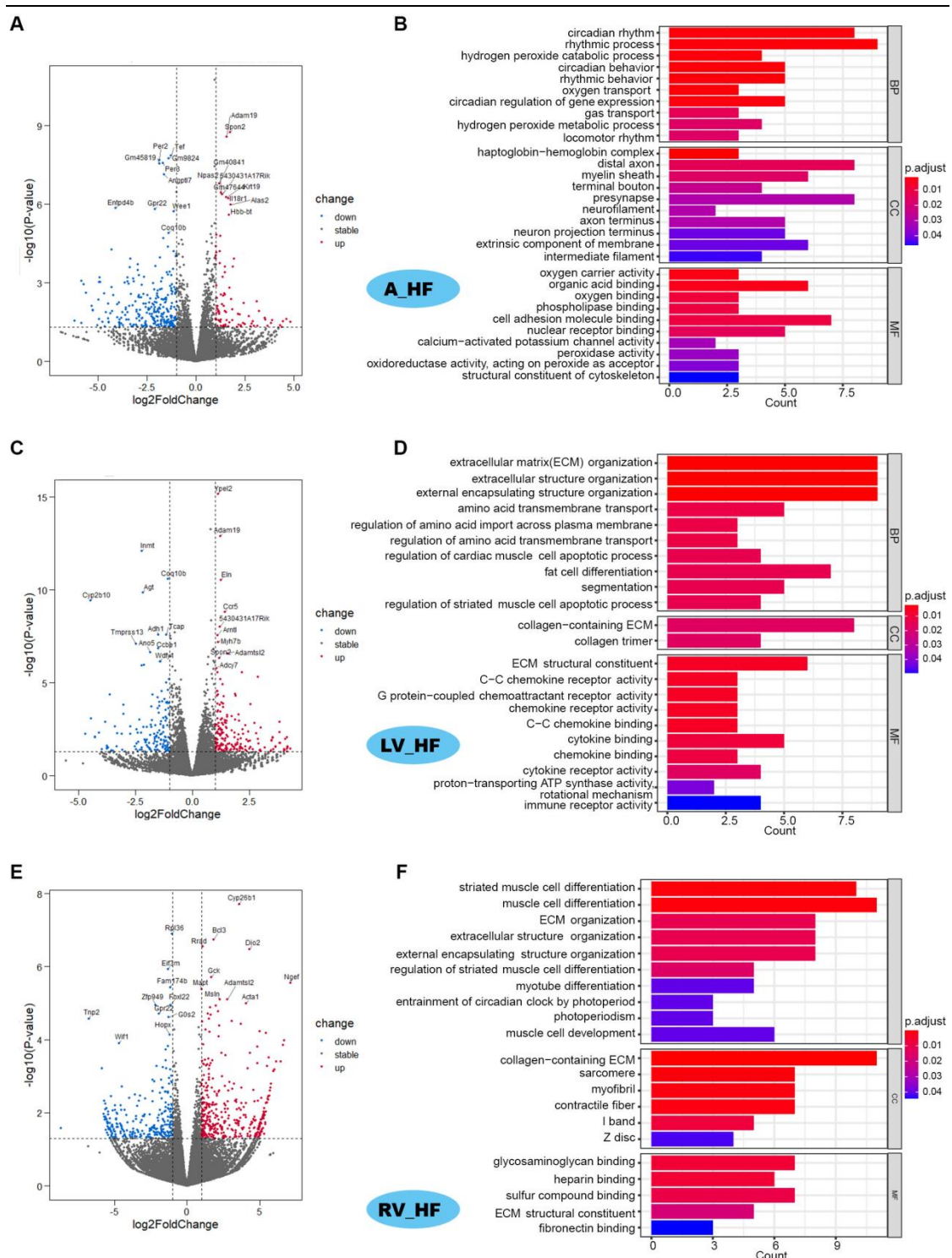


Figure S2 Analysis of differentially expressed genes (DEGs) in the same position under the HF-sham. Volcano plots of DEGs for in LA (A), LV (C), and RV (E), blue dots represent down-regulated DEGs; red dots represent up-regulated DEGs. The most significant 10 DEGs gene names are shown. The top 10 GO terms in BP, CC, and MF for DEGs in LA, LV, and RV are shown in the right part of the picture (B, D, F)

CH- cardiac hypertrophy, HF- heart failure, A- atrium (left), LV- left ventricle, RV- right ventricle, BP- biological process, CC- cellular component, MF- molecular function;

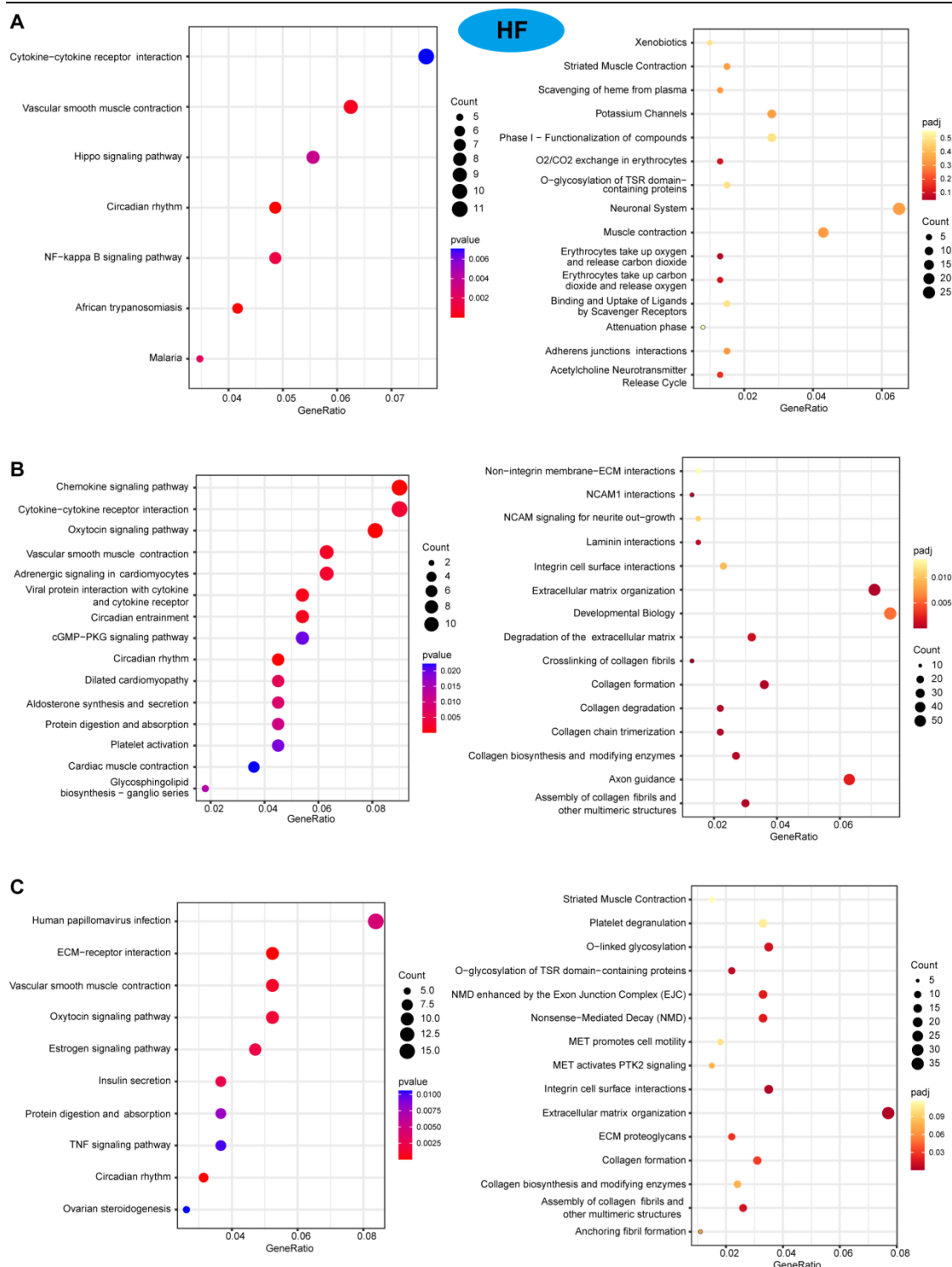
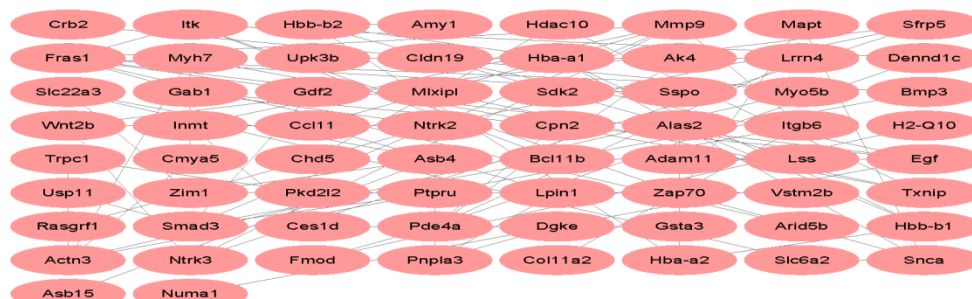
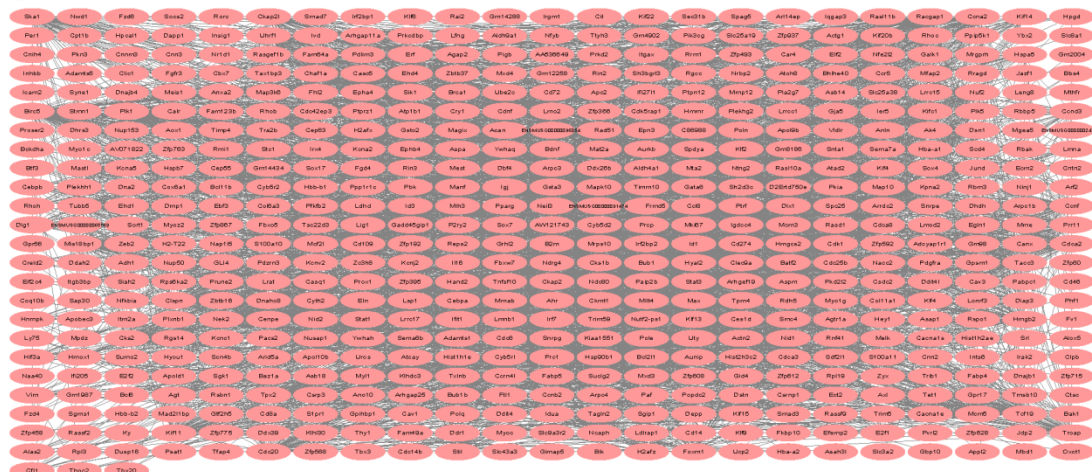


Figure S3 Statistics of pathway enrichment of DEG from the LA, LV, and RV of HF condition in mice. The KEGG enrichment pathway is on the left, and the Reactome enrichment pathway is on the right in the picture. (A) Enrichment pathway of the DEGs in the LA; (B) Enrichment pathway of the DEGs in LV; (C) Enrichment pathway of the DEGs in RV. The size of the point represents the number of DEGs
 CH- cardiac hypertrophy, HF- heart failure, A- atrium (left), LV- left ventricle, RV- right ventricle, BP- biological process, CC- cellular component, MF- molecular function; KEGG- Kyoto Encyclopedia of Genes and Genomes

A



B



C

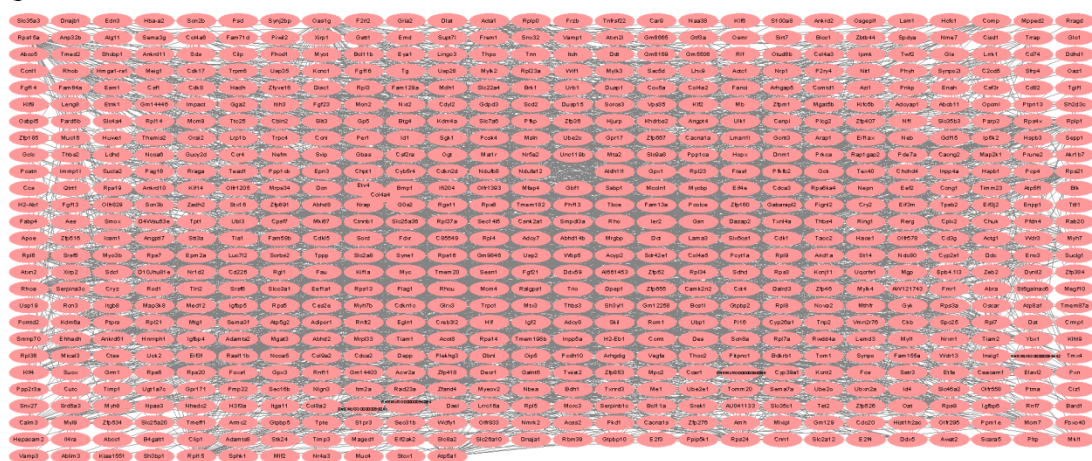


Figure S4 Constructing PPI network shared in CH and HF in LA, LV, and RV, respectively. (A) The PPI network of DEGs is shared in CH and HF in LA. (B) in LV. (C) in RV performed by the MCODE plugin in Cytoscape software

PPI- protein-protein interaction, CH- cardiac hypertrophy, HF- heart failure, DEGs- differentially expressed genes, LA- left atrium, LV- left ventricle, RV- right ventricle, MCODE- molecular complex detection.

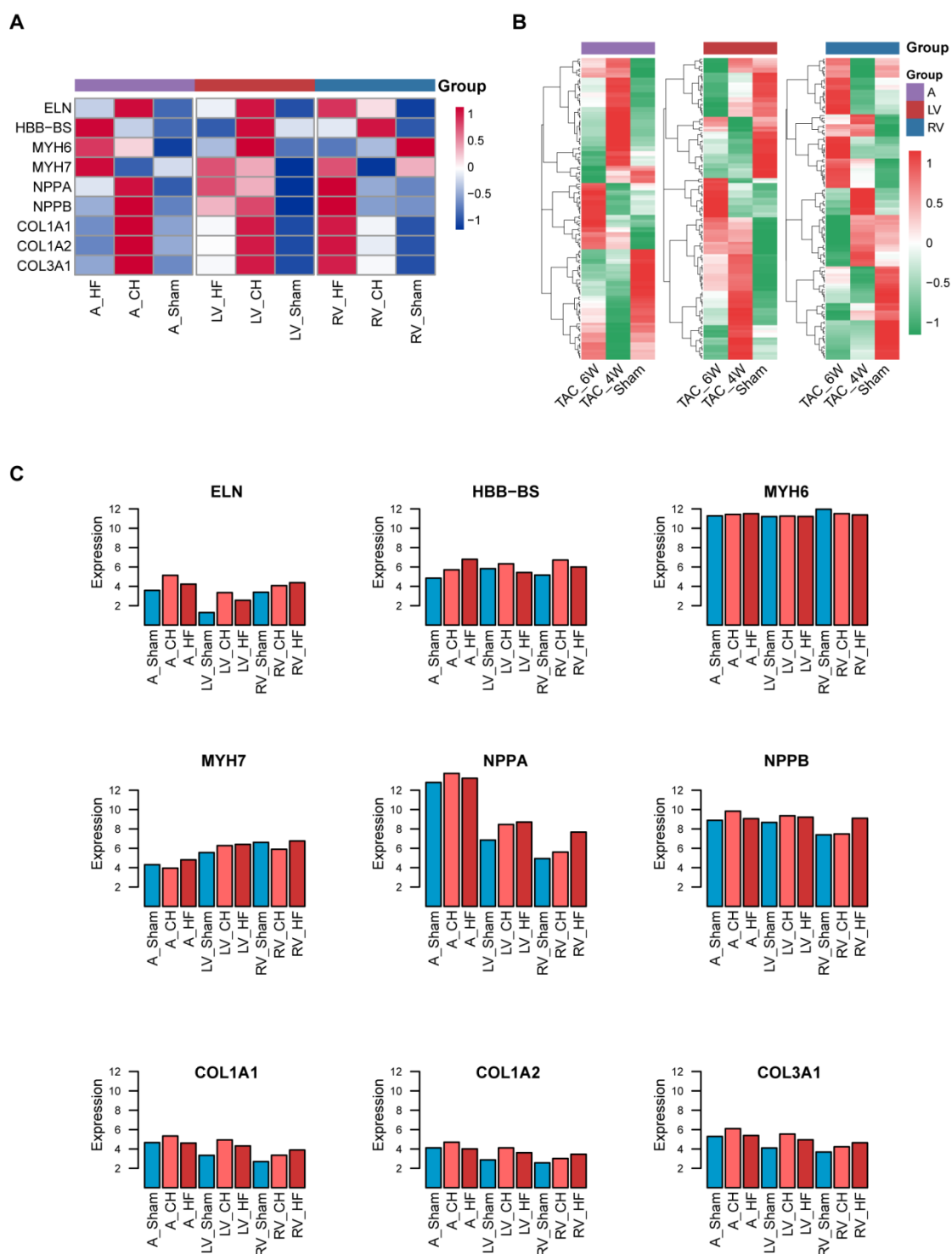


Figure S5 Expression of genes associated with cardiac hypertrophy and heart failure. (A) The heatmap of nine DEGs encoding cardiac myosin and natriuretic peptide hormones, fibrosis, and identified biomarkers in our study. (B) Heat maps of 123 genes may be known to be associated with hypertrophy and heart failure based on our gene expression dataset. (C) The expression of single DEG in LA, LV, and RV in different states
DEGs- differentially expressed genes, LA- left atrium, LV- left ventricule, RV- right ventricule