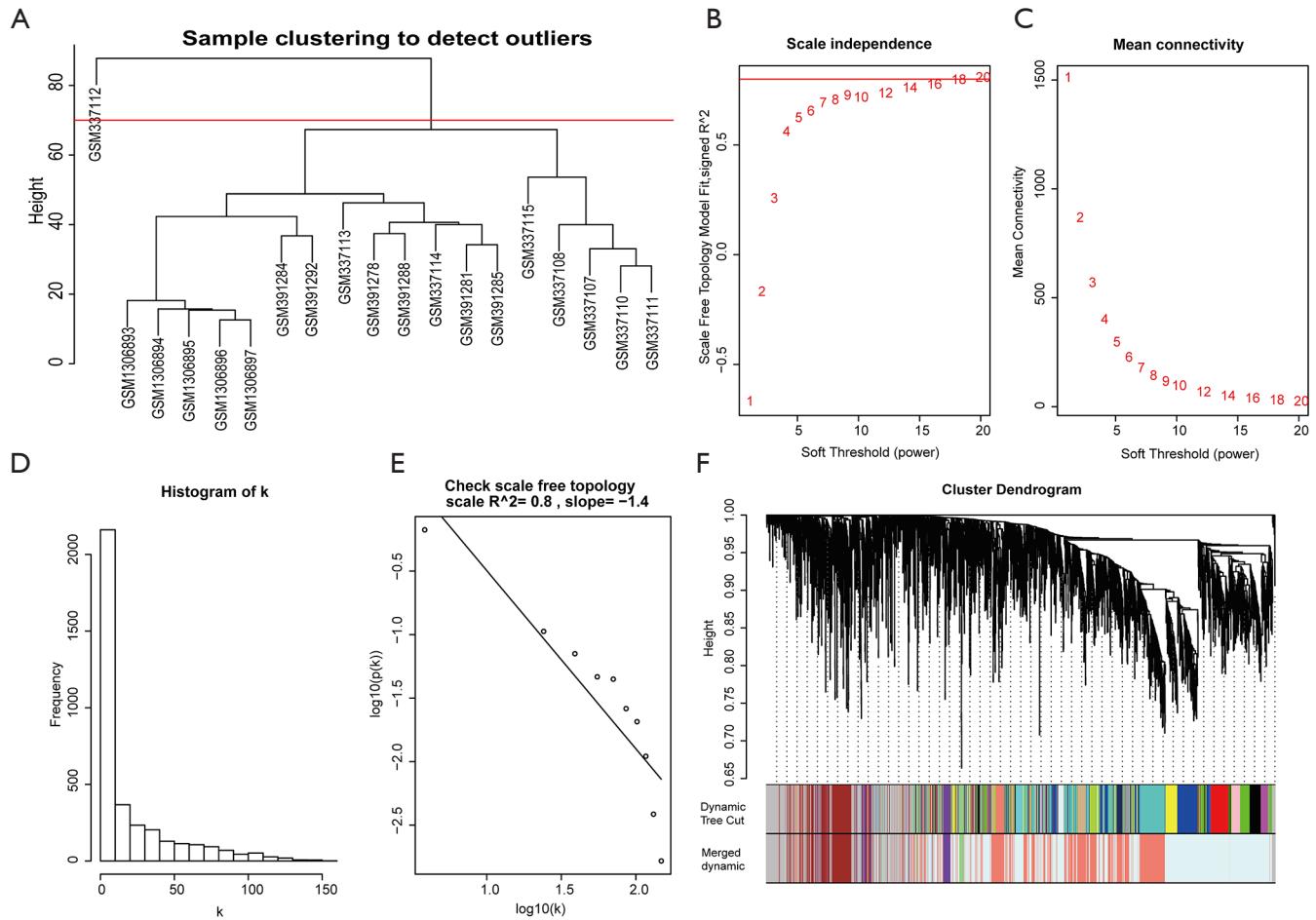


## Supplementary



**Figure S1** Construction of weighted gene co-expression network of unruptured IA samples. (A) Sample clustering detected 1 outliers (GSM337112). (B,C) The cut-off was set to be 0.8 and  $\beta=18$  was chosen to be the soft-threshold power. (D,E) The histogram and the linear fitting plot showed that the co-expression network we constructed met the requirement of scale-free topology. (F) 5 modules were detected after merging modules generated by dynamic tree cut. IA, intracranial aneurysm.

**Table S1** KEGG pathways for gene modules of ruptured IA and unruptured IA

Ruptured IA	KEGG pathways	Count	PValue	Unruptured IA	KEGG pathways	Count	PValue
Blue				Brown			
hsa04142:Lysosome	40	1.75E-19		hsa04151:PI3K-Akt signaling pathway	15	0.015286	
hsa05152:Tuberculosis	37	2.92E-11		hsa04514:Cell adhesion molecules (CAMs)	13	4.64E-05	
hsa04145:Phagosome	31	2.14E-09		hsa04510:Focal adhesion	12	0.004471	
hsa04380:Osteoclast differentiation	30	3.22E-10		hsa04723:Retrograde endocannabinoid signaling	11	5.33E-05	
hsa04062:Chemokine signaling pathway	29	3.61E-06		hsa04724:Glutamatergic synapse	11	1.49E-04	
hsa04015:Rap1 signaling pathway	25	0.001318		hsa04014:Ras signaling pathway	11	0.022573	
hsa04060:Cytokine-cytokine receptor interaction	25	0.008707		hsa04728:Dopaminergic synapse	10	0.001599	
hsa05164:Influenza A	24	2.10E-04		hsa04713:Circadian entrainment	9	9.09E-04	
hsa05323:Rheumatoid arthritis	23	4.24E-09		hsa04727:GABAergic synapse	8	0.002165	
hsa05150:Staphylococcus aureus infection	22	7.99E-13		hsa05032:Morphine addiction	8	0.003192	
hsa05140:Leishmaniasis	22	3.29E-10		hsa05033:Nicotine addiction	7	1.81E-04	
hsa04650:Natural killer cell mediated cytotoxicity	22	7.45E-06		hsa04911:Insulin secretion	7	0.009415	
hsa04514:Cell adhesion molecules (CAMs)	22	7.89E-05		hsa04726:Serotonergic synapse	7	0.031154	
hsa04620:Toll-like receptor signaling pathway	21	2.87E-06		hsa04670:Leukocyte transendothelial migration	7	0.036154	
hsa05145:Toxoplasmosis	20	1.93E-05		hsa04730:Long-term depression	6	0.00885	
hsa04670:Leukocyte transendothelial migration	20	3.68E-05		hsa04976:BilB secretion	6	0.015641	
hsa04611:Platelet activation	20	2.01E-04		hsa04012:ErbB signaling pathway	6	0.038014	
hsa01130:Biosynthesis of antibiotics	20	0.044292		hsa04512:ECM-receptor interaction	6	0.038014	
hsa04662:B cell receptor signaling pathway	18	3.18E-07		hsa05020:Prion diseases	5	0.00564	
hsa05142:Chagas disease (American trypanosomiasis)	18	1.08E-04		hsa05014:Amyotrophic lateral sclerosis (ALS)	5	0.021554	
hsa04666:Fc gamma R-mediated phagocytosis	17	2.52E-05		hsa05214:Gioma	5	0.049708	
hsa05162:Measles	17	0.004935		hsa00340:Histidine metabolism	4	0.010932	
hsa05133:Perlussis	16	2.46E-05		hsa0260:Glycine, serine and threonine metabolism	4	0.049933	
hsa04660:T cell receptor signaling pathway	16	7.03E-04		Lightcyan			
hsa04664:Fc epsilon RI signaling pathway	15	3.26E-05		hsa05200:Pathways in cancer	61	2.13E-06	
hsa04640:Hematopoietic cell lineage	15	5.11E-04		hsa04151:PI3K-Akt signaling pathway	44	0.004626	
hsa01200:Carbon metabolism	15	0.006405		hsa05205:Proteoglycans in cancer	38	2.60E-06	
hsa05134:Legionellosis	14	1.05E-05		hsa04810:Regulation of actin cytoskeleton	37	2.10E-05	
hsa03320:PPAR signaling pathway	14	1.19E-04		hsa04510:Focal adhesion	34	1.80E-04	
hsa05132:Salmonella infection	14	0.001044		hsa04022:cGMP-PKG signaling pathway	33	1.67E-06	
hsa04066:HIF-1 signaling pathway	14	0.003973		hsa04390:Hippo signaling pathway	30	1.52E-05	
hsa05169:Epstein-Barr virus infection	14	0.027185		hsa04015:Rap1 signaling pathway	30	0.004558	
hsa05146:Viral myocarditis	13	9.48E-05		hsa01130:Biosynthesis of antibiotics	29	0.00963	
hsa04064:NF-kappa B signaling pathway	13	0.004781		hsa04550:Signaling pathways regulating pluripotency of stem cells	27	7.58E-05	
hsa05321:Inflammatory bowel disease (IBD)	12	0.001149		hsa04242:cAMP signaling pathway	27	0.013237	
hsa05148:Amoebiasis	12	0.047315		hsa04270:Vascular smooth muscle contraction	25	2.68E-05	
hsa04672:Intestinal immune network for IgA production	11	3.31E-04		hsa04261:Adrenergic signaling in cardiomyocytes	25	3.99E-04	
hsa00520:Amino sugar and nucleotide sugar metabolism	10	0.001702		hsa0230:Purine metabolism	24	0.020098	
hsa04621:NOD-like receptor signaling pathway	10	0.005067		hsa04310:Wnt signaling pathway	23	0.002229	
hsa00010:Glycolysis / Gluconeogenesis	10	0.016177		hsa05010:Alzheimer's disease	22	0.039192	
hsa04610:Complement and coagulation cascades	10	0.019342		hsa04611:Platelet activation	21	0.005249	
hsa01230:Biosynthesis of amino acids	10	0.024898		hsa04921:Oxytocin signaling pathway	21	0.023828	
hsa04612:Antigen processing and presentation	10	0.033951		hsa04932:Non-alcoholic fatty liver disease (NAFLD)	20	0.045518	
hsa05144:Malaria	9	0.007326		hsa04152:AMPK signaling pathway	19	0.013022	
hsa05332:Graft-versus-host disease	8	0.002652		hsa04713:Circadian entrainment	18	0.001976	
hsa05330:Allograft rejection	8	0.005214		hsa05146:Amoebiasis	18	0.006372	
hsa04940:Type I diabetes mellitus	8	0.010606		hsa05222:Small cell lung cancer	17	0.001538	
hsa00480:Glutathione metabolism	8	0.028953		hsa04114:Oocyte meiosis	17	0.021193	
hsa05320:Autoimmune thyroid disease	8	0.031826		hsa01200:Carbon metabolism	17	0.024691	
hsa05310:Asthma	7	0.007206		hsa04350:TGF-beta signaling pathway	16	0.003636	
hsa05340:Primary immunodeficiency	7	0.013381		hsa05414:Dilated cardiomyopathy	16	0.003636	
hsa00052:Galactose metabolism	6	0.029335		hsa04512:ECM-receptor interaction	16	0.005121	
hsa00511:Other glycan degradation	5	0.018893		hsa04066:HIF-1 signaling pathway	16	0.012695	
hsa00531:Glycosaminoglycan degradation	5	0.022844		hsa04916:Melanogenesis	16	0.01807	
hsa04614:Renin-angiotensin system	5	0.043381		hsa04070:Phosphatidylinositol signaling system	15	0.032	
hsa00603:Glycosphingolipid biosynthesis - globo series	4	0.045674		hsa04915:Estrogen signaling pathway	15	0.034539	
hsa00290:Valine, leucine and isoleucine biosynthesis	3	0.019254		hsa04924:Renin secretion	14	0.002047	
Cyan				hsa05410:Hypertrophic cardiomyopathy (HCM)	14	0.011718	
hsa05200:Pathways in cancer	17	3.05E-04		hsa04970:Salivary secretion	14	0.025141	
hsa05166:HTLV-I infection	11	0.005568		hsa04666:Fc gamma R-mediated phagocytosis	13	0.044988	
hsa04151:PI3K-Akt signaling pathway	11	0.039259		hsa05412:Arrhythmogenic right ventricular cardiomyopathy (ARVC)	12	0.021728	
hsa05205:Proteoglycans in cancer	9	0.011744		hsa04971:Gastric acid secretion	12	0.038381	
hsa04510:Focal adhesion	9	0.013855		hsa05217:Basal cell carcinoma	11	0.012622	
hsa04668:TNF signaling pathway	8	0.001247		hsa04962:Vasopressin-regulated water reabsorption	10	0.009238	
hsa04310:Wnt signaling pathway	8	0.005254		hsa0280:Valine, leucine and isoleucine degradation	10	0.014206	
hsa05202:Transcriptional misregulation in cancer	8	0.014295		hsa04923:Regulation of lipolysis in adipocytes	10	0.040653	
hsa04978:Mineral absorption	7	5.10E-05		hsa00270:Cysteine and methionine metabolism	8	0.035282	
hsa04550:Signaling pathways regulating pluripotency of stem cells	7	0.020677		hsa00640:Propionate metabolism	7	0.025374	
hsa05222:Small cell lung cancer	6	0.009778		hsa04710:Circadian rhythm	7	0.040145	
hsa04974:Protein digestion and absorption	6	0.011264		Lightgreen			
hsa04916:Melanogenesis	6	0.01874		hsa04978:Mineral absorption	7	2.55E-08	
hsa05217:Basal cell carcinoma	5	0.009332		hsa00010:Glycolysis / Gluconeogenesis	5	1.86E-04	
hsa04640:Hematopoietic cell lineage	5	0.044799		hsa01130:Biosynthesis of antibiotics	5	0.012817	
hsa05323:Rheumatoid arthritis	5	0.046396		hsa01230:Biosynthesis of amino acids	4	0.003645	
Green				hsa01200:Carbon metabolism	4	0.012692	
hsa04141:Protein processing in endoplasmic reticulum	17	3.89E-07		hsa04068:FoxO signaling pathway	4	0.019992	
hsa04151:PI3K-Akt signaling pathway	15	0.012926		Purple			
hsa04510:Focal adhesion	14	3.54E-04		hsa04510:Focal adhesion	14	1.43E-09	
hsa04512:ECM-receptor interaction	11	1.19E-05		hsa04512:ECM-receptor interaction	10	8.92E-09	
hsa05205:Proteoglycans in cancer	10	0.024122		hsa04151:PI3K-Akt signaling pathway	10	7.29E-04	
hsa01130:Biosynthesis of antibiotics	10	0.03342		hsa04670:Leukocyte transendothelial migration	7	1.67E-04	
hsa03050:Proteasome	9	2.84E-06		hsa04810:Regulation of actin cytoskeleton	7	0.003903	
hsa05222:Small cell lung cancer	8	0.001924		hsa05222:Small cell lung cancer	6	3.50E-04	
hsa04145:Phagosome	8	0.037561		hsa05146:Amoebiasis	6	9.65E-04	
hsa04974:Protein digestion and absorption	7	0.010067		hsa04514:Cell adhesion molecules (CAMs)	6	0.003508	
hsa05146:Amoebiasis	7	0.023388		hsa05205:Proteoglycans in cancer	6	0.014498	
hsa05169:Epstein-Barr virus infection	7	0.042388		hsa05100:Bacterial invasion of epithelial cells	5	0.002373	
hsa00510-N-Glycan biosynthesis	6	0.003409		hsa05410:Hyperthrophic cardiomyopathy (HCM)	5	0.002373	
hsa05130:Pathogenic Escherichia coli infection	6</						

**Table S2** Hub genes of each gene module

Module	Gene ID	Gene symbol	Official full name
Ruptured IA			
Blue	9056	SLC7A7	Solute carrier family 7 member 7
Cyan	9945	GFPT2	Glutamine-fructose-6-phosphate transaminase 2
Green	378	ARF4	ADP ribosylation factor 4
Green yellow	4118	MAL	Mal, T cell differentiation protein
Lightcyan	257240	KLHL34	Kelch like family member 34
Midnightblue	5332	PLCB4	Phospholipase C beta 4
Purple	5592	PRKG1	Protein kinase cGMP-dependent 1
Salmon	8875	VNN2	Vanin 2
Unruptured IA			
Brown	590	BCHE	Butyrylcholinesterase
Lightcyan	4012	LNPEP	Leucyl and cysteinyl aminopeptidase
Lightgreen	4495	MT1G	Metallothionein 1G
Purple	91663	MYADM	Myeloid associated differentiation marker
Salmon	963	CD53	CD53 molecule

IA, intracranial aneurysm.

**Table S3** Validation of key genes using dataset GSE122897

Gene symbol	t	P value	P value summary
CXCR4	2.029	0.0492	*
AQP9	2.374	0.0225	*
SLA	2.712	0.0098	**
MPP1	4.044	0.0002	***
PDZRN3	2.639	0.0118	*
ANGPT1	3.017	0.0044	**
FPR1	2.430	0.0197	*
ANPEP	3.111	0.0034	**
SRGN	2.417	0.0203	*
C15orf48	3.314	0.0020	**
NCF2	2.732	0.0093	**
C1orf162	3.024	0.0043	**
TCIRG1	3.018	0.0044	**
FGR	2.259	0.0294	*

\*, P&lt;0.05; \*\*, P&lt;0.01; \*\*\*, P&lt;0.001.

**Table S4** ROC analysis of key genes

Gene symbol	AUC	SE	P value
<i>CXCR4</i>	0.703	0.082	0.024
<i>AQP9</i>	0.717	0.082	0.016
<i>SLA</i>	0.726	0.080	0.012
<i>MPP1</i>	0.821	0.069	0.000
<i>PDZRN3</i>	0.692	0.085	0.034
<i>ANGPT1</i>	0.766	0.077	0.003
<i>FPR1</i>	0.696	0.081	0.030
<i>ANPEP</i>	0.737	0.078	0.009
<i>SRGN</i>	0.714	0.082	0.017
<i>C15orf48</i>	0.764	0.076	0.003
<i>NCF2</i>	0.728	0.078	0.011
<i>C1orf162</i>	0.757	0.081	0.004
<i>TCIRG1</i>	0.766	0.077	0.003
<i>FGR</i>	0.712	0.080	0.019

ROC, receiver operating characteristic; AUC, area under curve; SE, standard error.