Supplementary figures and tables of "Reconstruction of human protein-coding gene functional association network based on machine learning"

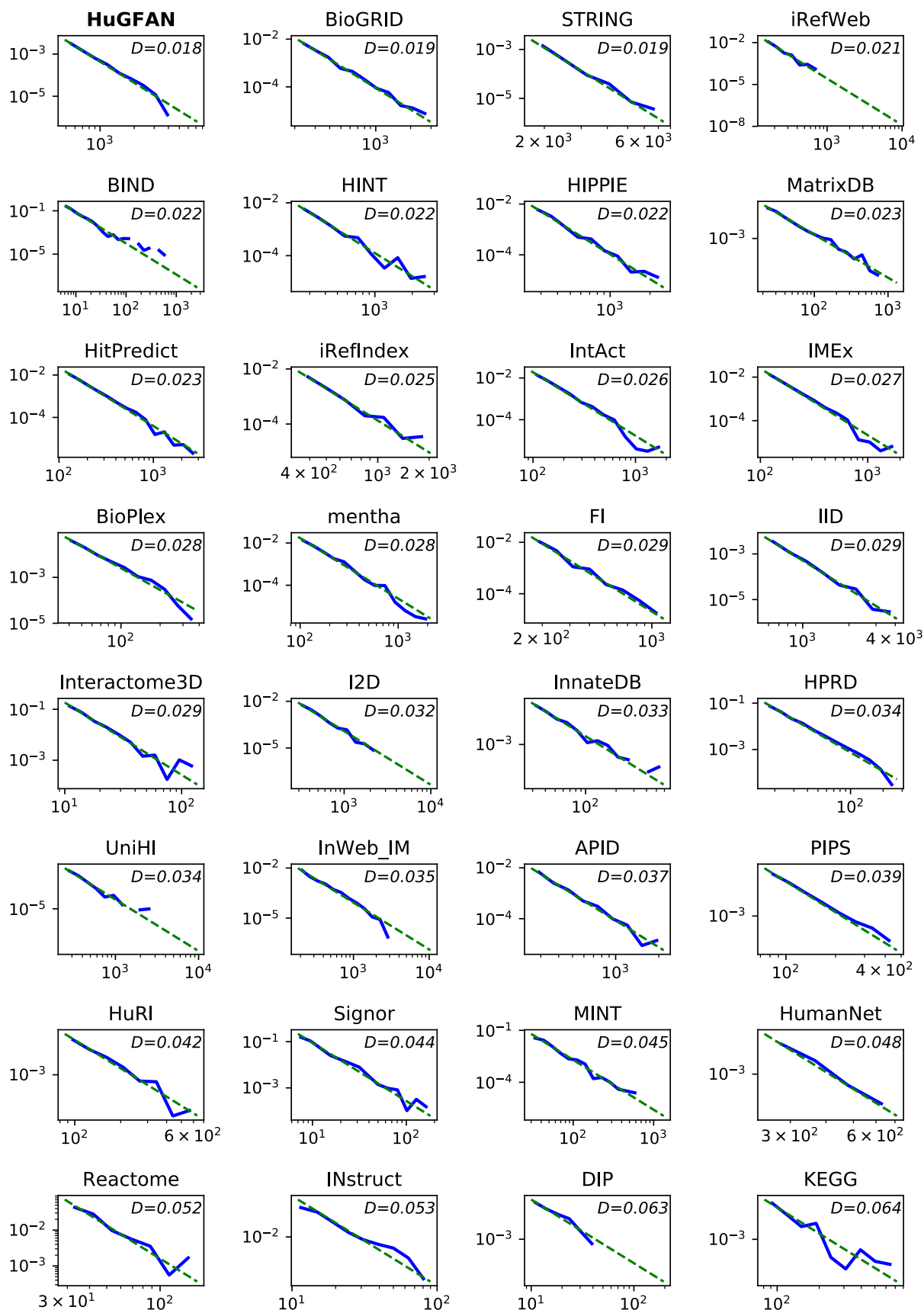HUANG Xiaotai*, Songwei Jia and Lin Gao

Figure S1: The power law fitting of HuGFAN and other networks. Ranked by Kolmogorov-Smirnov distance (D).
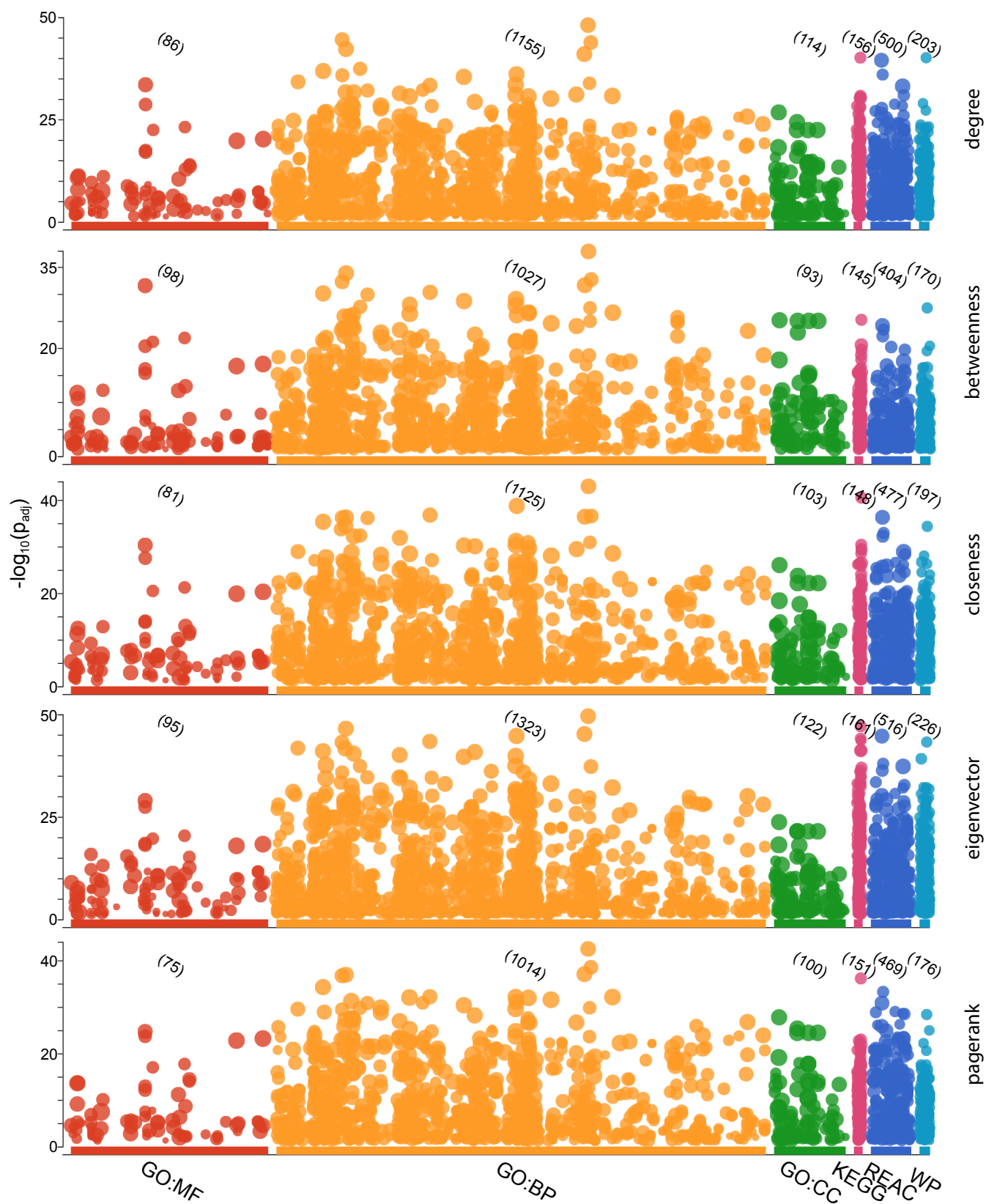
Figure S2: The enrichment results of GO terms and pathways for HuGFAN top 100 genes in rankings of the five centralities.

Table S1: Network topology metrics comparison between HuGFAN and other networks.

Networks	Avg. CC	Avg. Degree	Avg. Betweenness	Avg. Closeness	Avg. Eigenvector	Avg. Pagerank
STRING	0.203	594.173	9669.360	0.491	0.088	5.39E-05
HuGFAN	0.296	116.315	16968.672	0.380	0.031	4.91E-05
IID	0.243	106.690	15784.024	0.372	0.039	5.46E-05
InWeb_IM	0.317	74.086	14235.404	0.389	0.025	5.68E-05
HumanNet	0.287	58.590	18702.214	0.328	0.020	5.58E-05
HINT	0.150	51.588	16413.065	0.352	0.032	5.71E-05
HitPredict	0.118	47.435	16494.430	0.347	0.025	5.81E-05
BioGRID	0.129	45.907	17893.537	0.347	0.024	5.35E-05
iRefIndex	0.185	45.366	17677.007	0.334	0.020	5.74E-05
HIPPIE	0.133	42.254	17553.609	0.343	0.030	5.57E-05
UniHI	0.328	41.192	15226.191	0.375	0.014	5.65E-05
mentha	0.109	40.305	17454.641	0.334	0.026	5.80E-05
APID	0.125	39.441	17310.106	0.332	0.027	5.93E-05
FI	0.422	38.021	15772.794	0.309	0.023	7.39E-05
I2D	0.254	35.390	13429.346	0.386	0.014	6.11E-05
IntAct	0.098	30.455	18397.342	0.319	0.020	5.91E-05
IMEx	0.093	30.176	18276.813	0.320	0.020	5.91E-05
iRefWeb	0.254	20.678	13159.471	0.370	0.010	6.74E-05
PIPS	0.341	19.807	8636.076	0.285	0.029	1.55E-04
BioPlex	0.112	16.914	19757.916	0.265	0.012	7.17E-05
KEGG	0.271	16.909	9535.088	0.249	0.023	1.72E-04
HuRI	0.082	13.946	11867.599	0.280	0.024	1.12E-04
MatrixDB	0.137	13.647	14383.743	0.290	0.017	8.76E-05
Reactome	0.600	9.408	5334.450	0.350	0.019	2.19E-04
HPRD	0.136	7.761	14483.688	0.258	0.022	1.06E-04
MINT	0.154	6.853	11402.708	0.264	0.009	1.28E-04
Signor	0.105	5.345	6416.482	0.272	0.025	2.30E-04
InnateDB	0.185	4.759	8088.615	0.263	0.021	1.81E-04
Interactome3D	0.270	4.672	10387.688	0.296	0.014	1.70E-04
BIND	0.288	4.571	7772.695	0.324	0.013	1.41E-04
DIP	0.283	3.269	4946.810	0.290	0.008	3.29E-04
INstruct	0.197	3.260	2876.147	0.389	0.014	3.82E-04

Note: Abbreviations used in the table are: **CC**: Clustering Coefficient, **Avg.**: average.

Table S2: Prediction performance comparison between HuGFAN and other networks with DriverNet.

Networks	GBM			MT			OV			TN			Avg.
	p-value	ACC	Rank	p-value	ACC	Rank	p-value	ACC	Rank	p-value	ACC	Rank	Rank
HuGFAN	0.001252	0.788	**1**	0.049148	0.779	**1**	0.000844	0.808	**1**	0.000859	0.911	2	**1.25**
IID	0.003487	0.688	2	0.123308	0.759	2	0.127765	0.749	8	0.024179	0.898	4	4
STRING	0.028464	0.679	3	0.305066	0.358	14	0.005747	0.781	2	0.025183	0.903	3	5.5
InWeb_IM	0.010117	0.654	5	0.571521	0.719	3	0.040270	0.764	3	0.364030	0.893	12	5.75
HINT	0.107288	0.627	9	0.519966	0.700	4	0.128572	0.702	14	0.057298	0.908	5	8
FI	0.013890	0.661	4	0.830629	0.640	9	0.012889	0.744	4	0.432827	0.863	17	8.5
BioGRID	0.027654	0.617	6	0.845064	0.509	10	0.357777	0.694	15	0.075735	0.897	8	9.75
HitPredict	0.095833	0.609	10	0.741535	0.672	5	0.376187	0.677	17	0.097226	0.890	13	11.25
HumanNet	0.179893	0.510	16	1	0.094	23	0.003997	0.723	5	0.000357	0.912	**1**	11.25
APID	0.088798	0.511	15	0.935337	0.643	8	0.597470	0.673	18	0.196577	0.904	6	11.75
UniHI	0.142188	0.593	12	0.611380	0.177	17	0.484708	0.729	11	0.302609	0.896	9	12.25
I2D	0.186039	0.523	14	0.644045	0.200	16	0.485799	0.738	9	0.068290	0.882	15	13.5
iRefIndex	0.185131	0.600	11	0.728953	0.148	18	0.054469	0.736	10	0.199221	0.874	16	13.75
HIPPIE	0.507726	0.469	19	0.800213	0.670	6	0.384832	0.657	21	0.094043	0.895	10	14
INstruct	1	0.667	8	1	0.250	15	0.500000	0.750	7	1	0.500	26	14
BIND	1	0.500	17	1	0.667	7	1	0.692	16	0.333333	0.833	19	14.75
mentha	0.153352	0.488	18	0.621680	0.429	12	0.226005	0.662	19	0.079393	0.894	11	15
DIP	1	0.667	8	1	0.500	11	0.700000	0.600	22	1	0.750	23	16
iRefWeb	0.211888	0.500	17	1	0.097	22	0.339559	0.724	12	0.212721	0.861	18	17.25
IntAct	0.456140	0.421	20	1	0.139	20	0.408214	0.571	23	0.114678	0.903	7	17.5
KEGG	1	0.250	25	0.752632	0.381	13	0.093255	0.707	13	0.075475	0.789	21	18
PIPS	1	0.364	22	1	0.125	21	0.033214	0.717	6	0.302150	0.692	25	18.5
IMEx	1	0.350	23	1	0.147	19	0.867323	0.478	25	0.165607	0.888	14	20.25
Interactome3D	1	0.714	7	1	0	25	1	0.286	29	0.109091	0.818	20	20.25
MatrixDB	1	0.400	21	1	0.091	24	0.577421	0.659	20	0.437835	0.833	19	21
HuRI	1	0.571	13	1	0	25	0.821429	0.5	24	0.450000	0.750	23	21.25
HPRD	1	0.500	17	1	0	25	0.761206	0.435	27	1	0.773	22	22.75
MINT	1	0.500	17	1	0	25	1	0.231	30	1	0.700	24	24
BioPlex	1	0.143	26	1	0.200	16	0.828431	0.333	28	0.961539	0.231	30	25
Signor	1	0.400	21	1	0	25	0.803030	0.455	26	1	0.286	29	25.25
Reactome	1	0.333	24	1	0	25	1	0.111	32	0.892857	0.375	28	27.25
InnateDB	1	0	27	1	0	25	1	0.2	31	1	0.400	27	27.5

Note: Abbreviations used in the table are: **GBM**: glioblastoma, **TN**: triple negative breast cancers, **MT**: breast tumors, **OV**: serous ovarian cancers.

Table S3: Prediction performance comparison between HuGFAN and other networks with HotNet2.

Networks	# Subnetworks	Max_sub	# Genes	# CGC genes	Pan-cancer enrich.		KEGG cancer enrich.		Avg. Rank
					# Pan-cancer_sub	Rank	# Cancer_path	Rank	
HuGFAN	29	246	389	94	13	**1**	17	**1**	**1**
KEGG	36	111	264	86	6	6	12	3	4.5
HPRD	33	19	140	48	10	2	7	7	4.5
DIP	17	25	103	52	5	7	12	3	5
BioGRID	29	95	200	58	8	4	8	6	5
BioPlex	118	27	541	96	4	8	12	3	5.5
Interactome3D	53	73	333	98	5	7	10	4	5.5
HitPredict	31	76	191	51	8	4	7	7	5.5
InWeb_IM	12	126	170	52	8	4	7	7	5.5
INstruct	18	60	154	53	2	10	13	2	6
Reactome	23	45	139	54	5	7	9	5	6
HINT	27	76	178	47	8	4	6	8	6
UniHI	20	80	153	47	9	3	5	9	6
STRING	136	284	848	93	7	5	6	8	6.5
MatrixDB	28	32	158	48	4	8	8	6	7
mentha	29	82	185	49	8	4	4	10	7
PIPS	67	32	325	57	0	12	12	3	7.5
IntAct	20	77	144	47	6	6	5	9	7.5
HIPPIE	25	96	192	53	7	5	4	10	7.5
FI	88	120	488	76	8	4	3	11	7.5
HuRI	134	46	758	91	3	9	7	7	8
APID	31	81	196	48	7	5	3	11	8
iRefWeb	22	43	129	44	7	5	3	11	8
I2D	30	68	189	47	7	5	2	12	8.5
IMEx	7	24	51	37	7	5	2	12	8.5
HumanNet	21	66	152	42	5	7	3	11	9
IID	39	111	295	49	6	6	2	12	9
iRefIndex	28	96	211	55	6	6	2	12	9
MINT	15	42	89	41	7	5	1	13	9
BIND	18	27	114	52	2	10	5	9	9.5
Signor	16	65	126	57	1	11	4	10	10.5
InnateDB	16	34	124	49	1	11	1	13	12

Note: Abbreviations used in the table are: **# (number sign)**: number of, **Max_sub**: maximum size of subnetworks, **enrich.**: enrichment, **Pan-cancer_sub**: the identified subnetworks in [1], **Cancer_path**: KEGG pathways in cancer.

References

[1] Mark DM Leiserson, Fabio Vandin, Hsin-Ta Wu, Jason R Dobson, Jonathan V Eldridge, Jacob L Thomas, Alexandra Papoutsaki, Younhun Kim, Beifang Niu, Michael McLellan, et al. Pan-cancer network analysis identifies combinations of rare somatic mutations across pathways and protein complexes. *Nature genetics*, 47(2):106–114, 2015.