

(a) Protein complexes

Network	SRCC _{degree}	SRCC _{degree/size}	SRCC _{degree·size}
Direct	0.3112 (3e-10)	0.2795 (2e-08)	0.1974 (9e-05)
Pull-down	0.2933 (4e-09)	0.2294 (5e-06)	0.1753 (5e-04)
Full	0.3093 (4e-10)	0.2288 (5e-06)	0.1924 (1e-04)

(b) Filtered biological processes

Network	SRCC _{degree}	SRCC _{degree/size}	SRCC _{degree·size}
Direct	0.3325 (2e-11)	0.2186 (1e-05)	0.2220 (1e-05)
Pull-down	0.3786 (9e-15)	0.2608 (2e-07)	0.2734 (4e-08)
Full	0.3475 (2e-12)	0.1068 (3e-02)	0.2384 (2e-06)

Table S 5. Correlation between cross-talk (CT) degree and binary module essentiality after removing functionally similar cross-talks for (a) protein complexes and (b) filtered biological processes. We removed cross-talks between modules that are annotated with a shared general biological process (see main body of the paper). In this restricted module-level network, **SRCC_{degree}** gives the SRCC between the cross-talk degree and essentiality. **SRCC_{degree/size}** gives the SRCC between the normalized cross-talk degree (i.e., cross-talk degree of the module divided by the module size) and essentiality. **SRCC_{degree·size}** gives the partial SRCC between cross-talk degree and essentiality when controlling for the module size. *P*-values for the SRCCs are shown within parentheses. A module is considered essential if has at least one essential protein, and not essential otherwise.