



Multiplex Biological Networks

Communities



I2M

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BioNetVisa WS

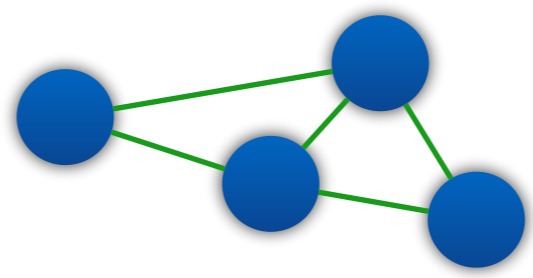
25/11/2015

Singapore

Systems Biology



Genotypes



**-Omics
Networks**

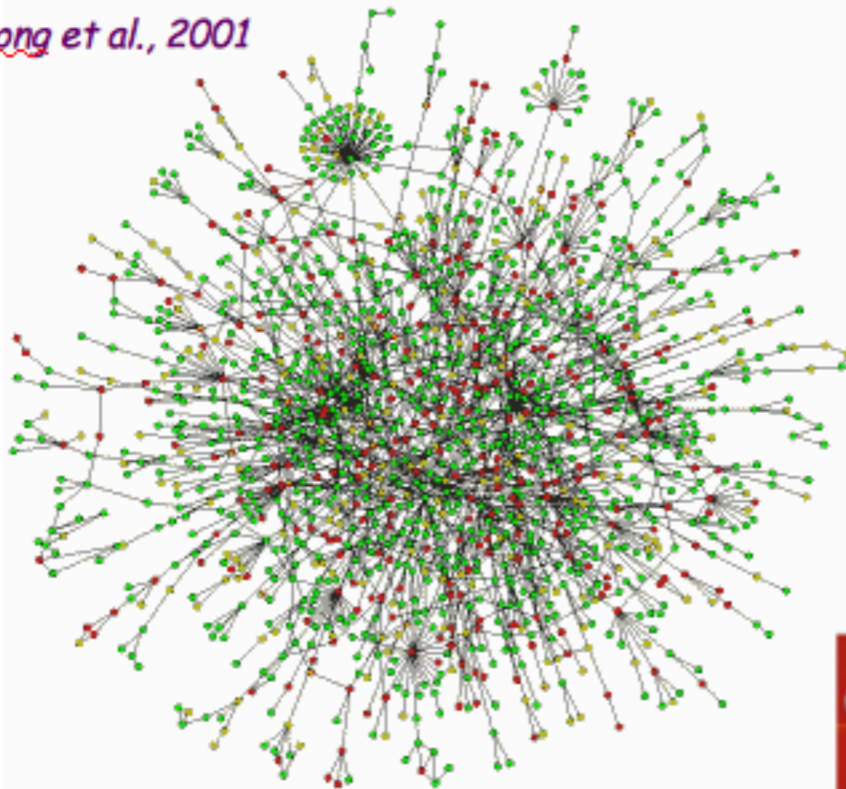


Phenotypes

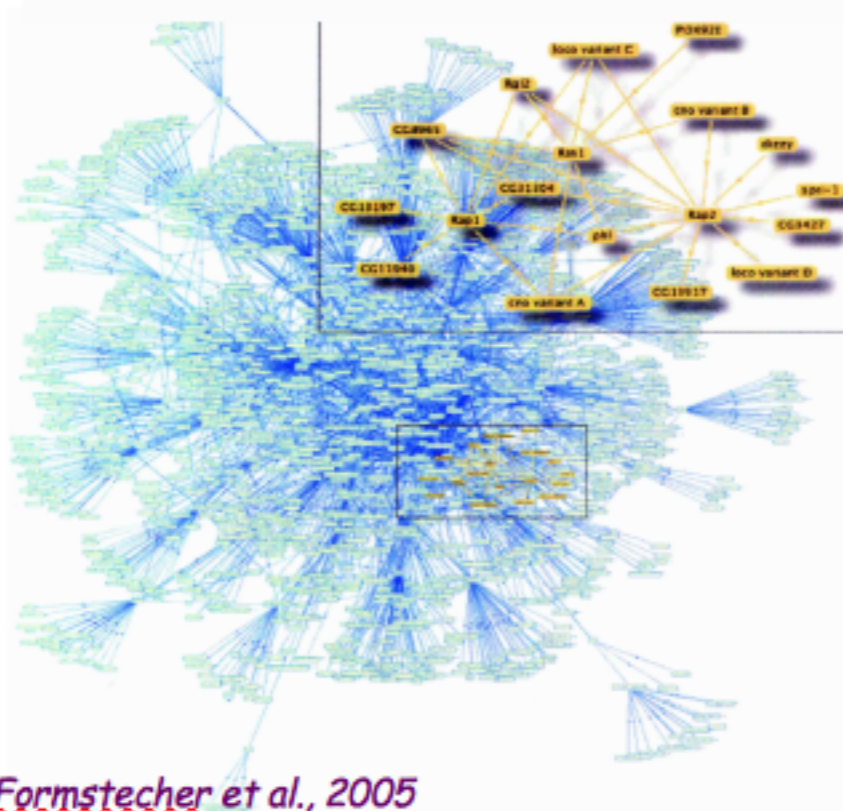
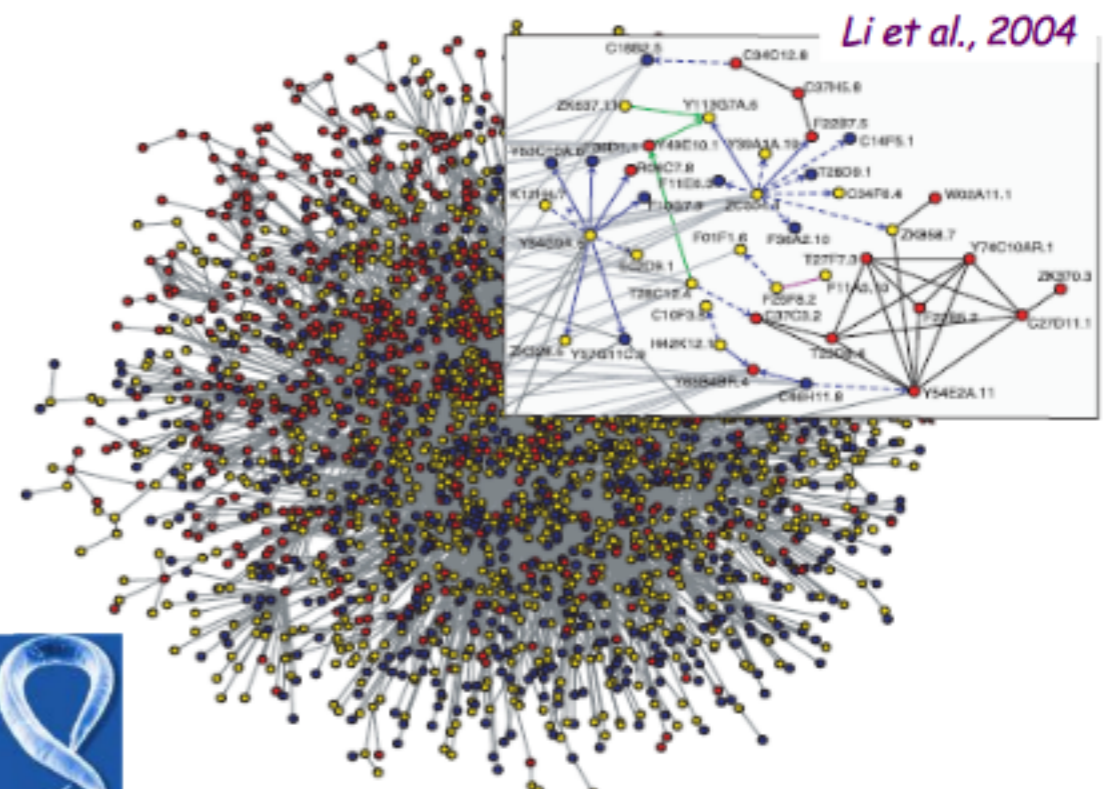


Large-scale Protein-protein Interaction Networks

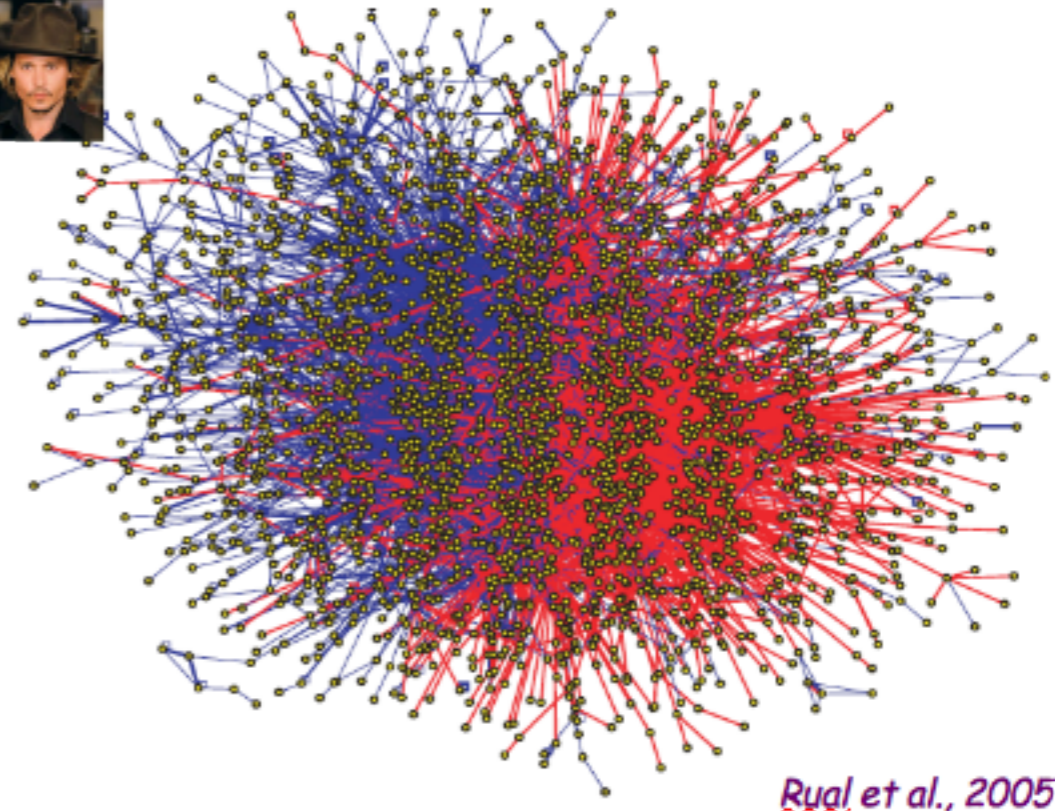
Jeong et al., 2001



Li et al., 2004



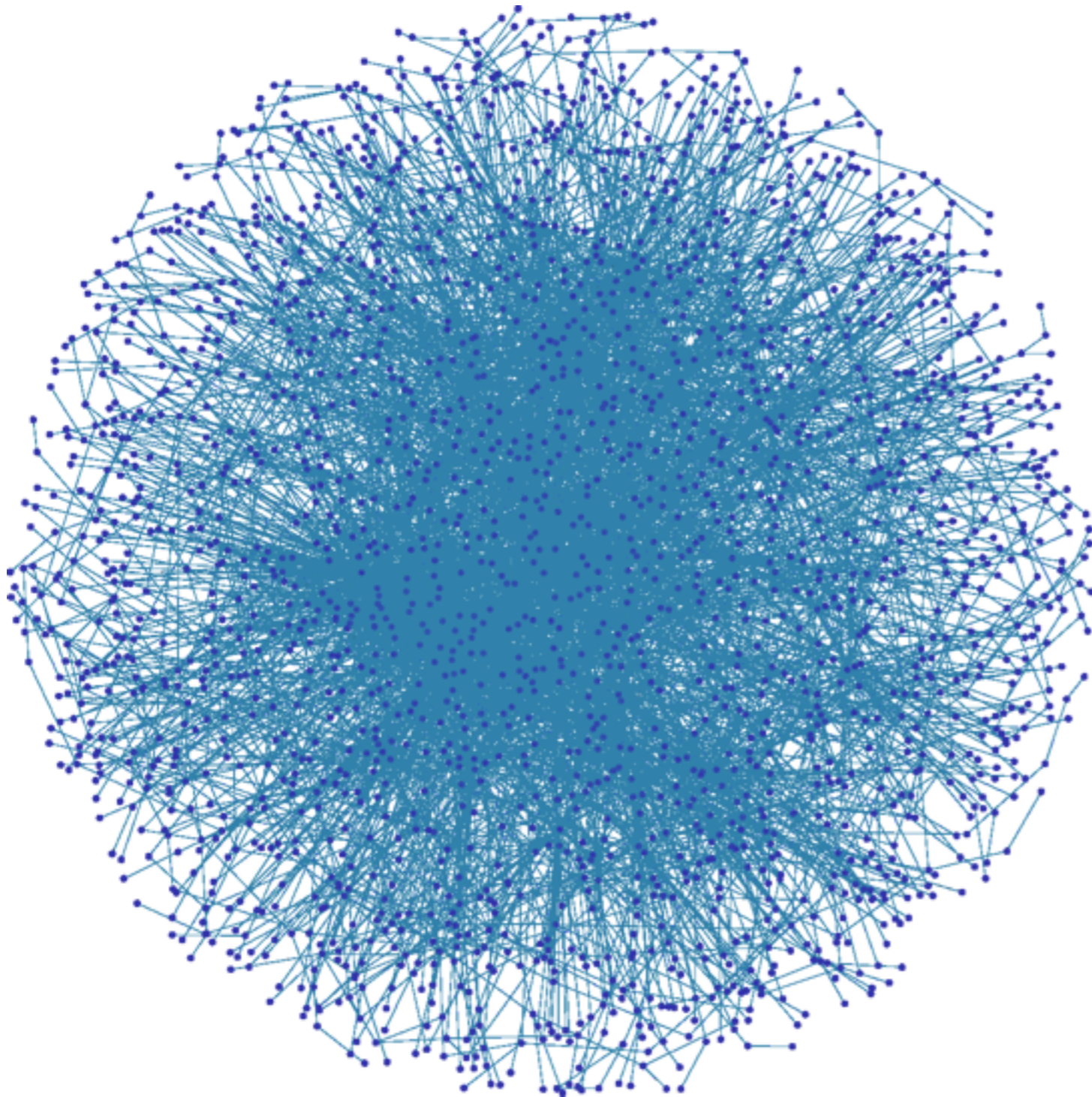
Formstecher et al., 2005



Rual et al., 2005



Interactomes

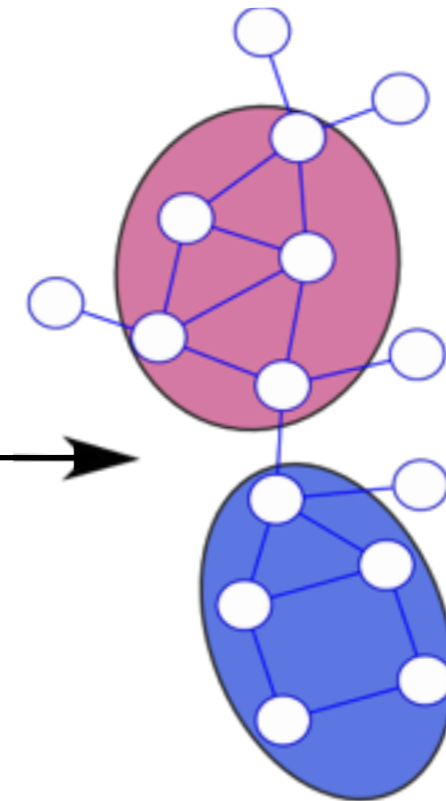
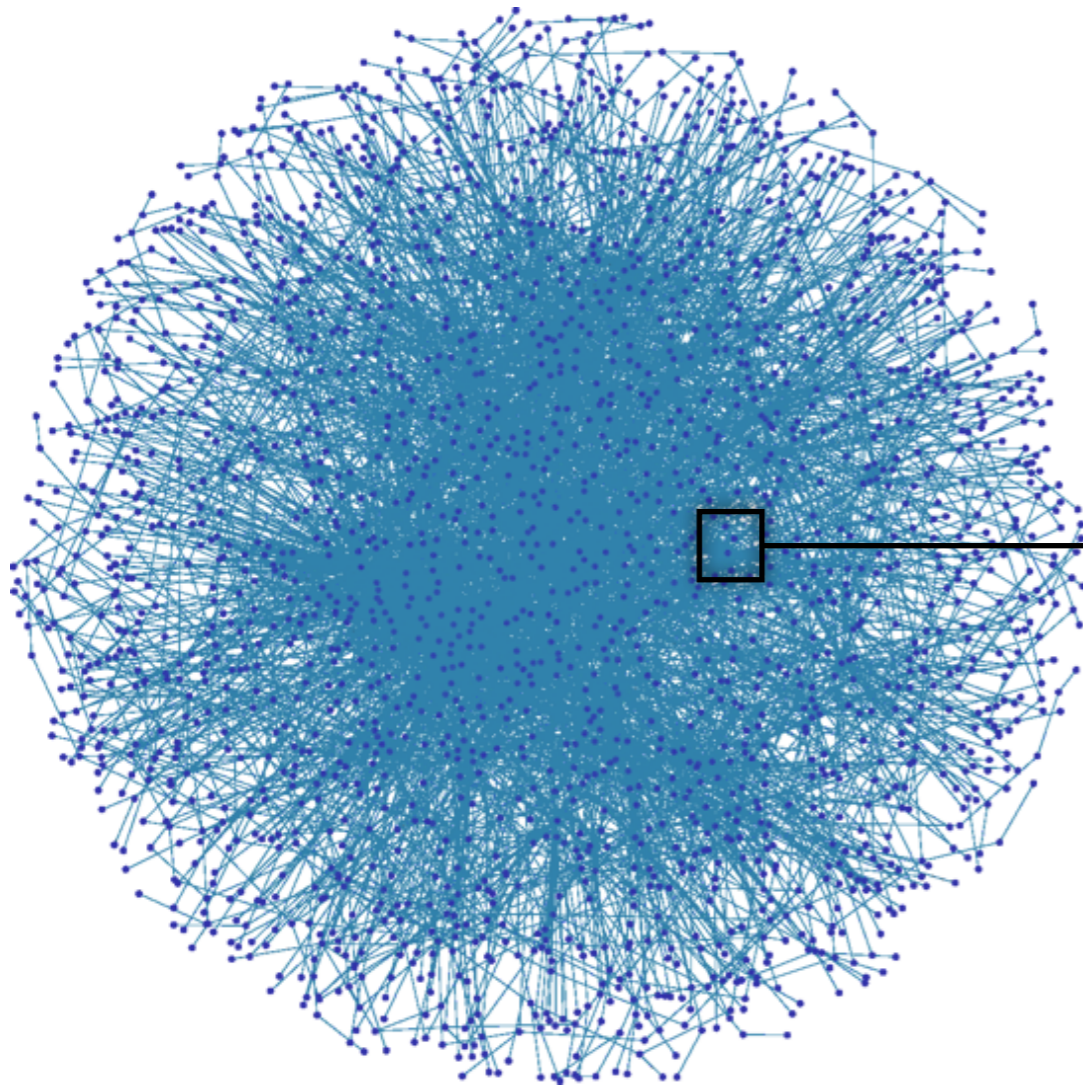


How to extract relevant biological information ?



Identification of communities

Classes / Clusters / Modules



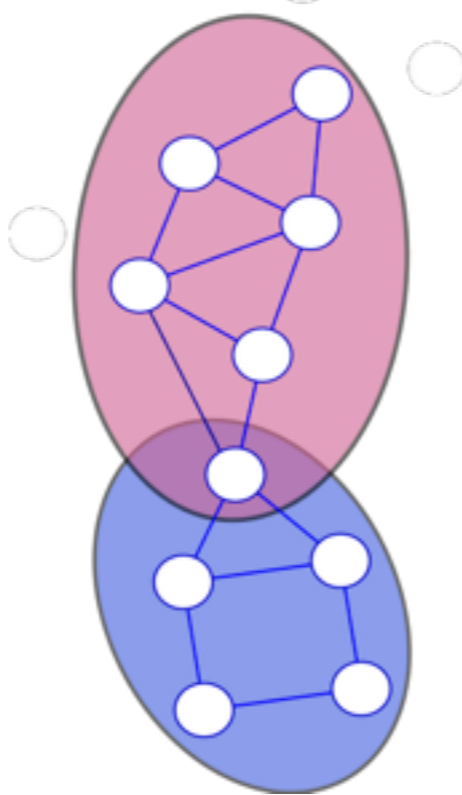
Group of tightly linked genes/proteins



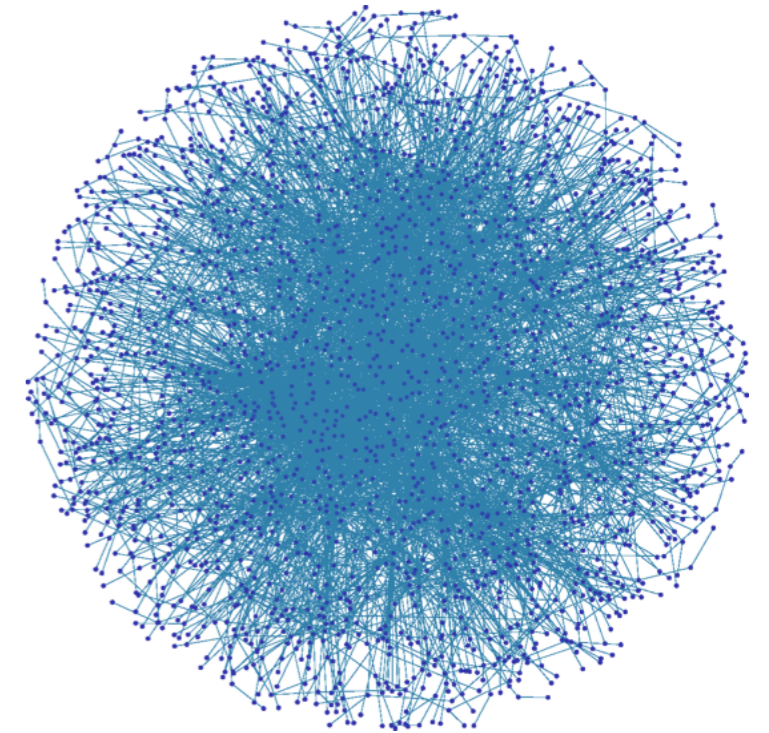
Protein cellular functions



Protein - Structures



Functional modules



Interaction Networks

Precision

Comprehensiveness

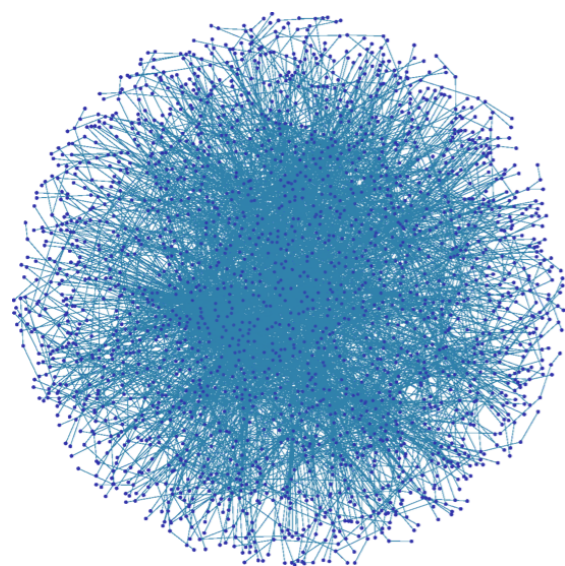
Inspired from P. Aloy, ECCB 2014

- Topology : Density, **Modularity**
- Integration of external data
- Overlapping / Disjoint



Multiple networks to depict protein cellular functions

PPI (12110 / 60669)



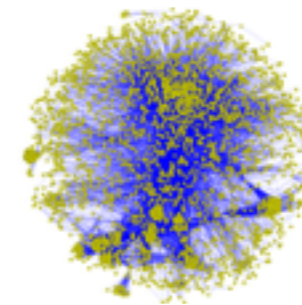
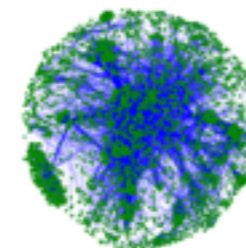
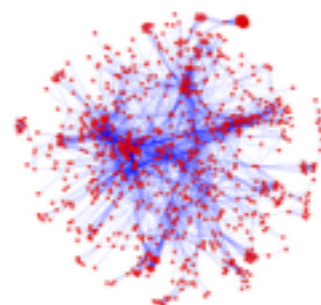
Pathway-related (8839 / 166761)

Spike

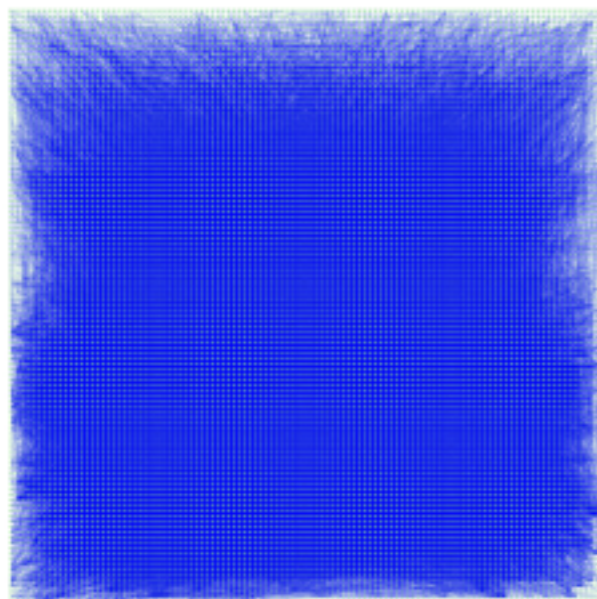
Biocarta

Kegg

Nci

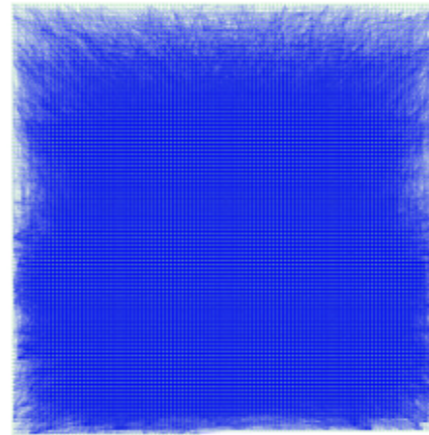
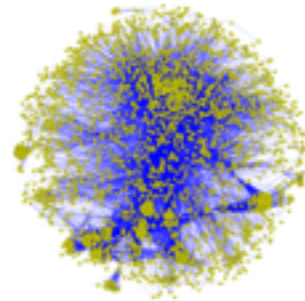
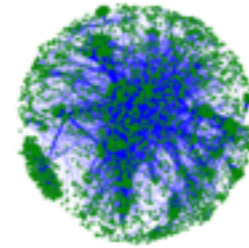
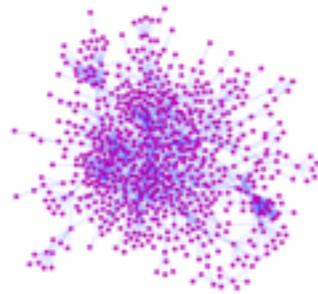
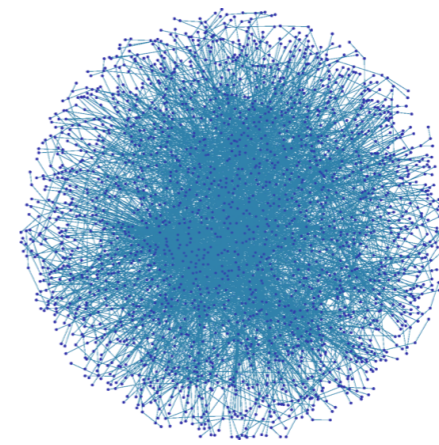
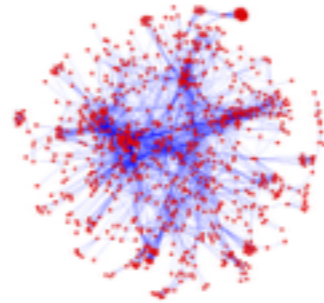


Co-Expression (9912 / 1107547)



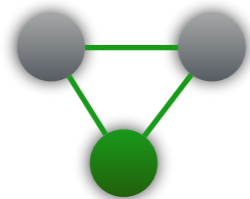
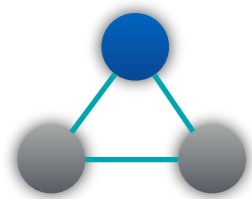
CORUM complexes (2528 / 36762)



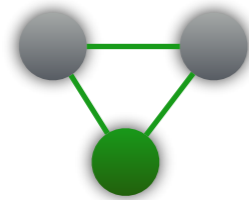
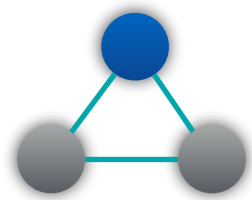
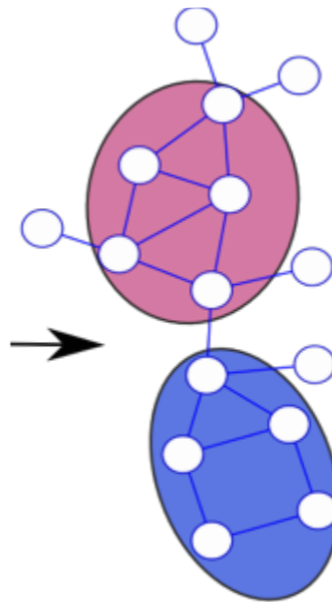
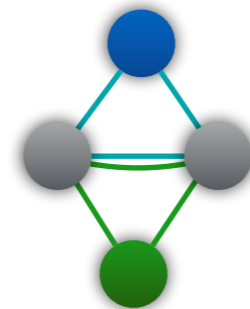


- **Does the combination of networks improves the detection of communities?**
- **How to combine multiple networks to identify communities?**

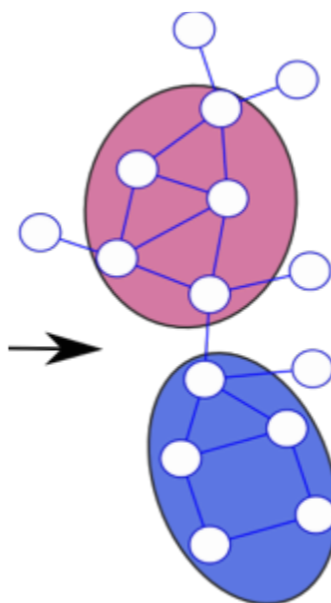
Network aggregation



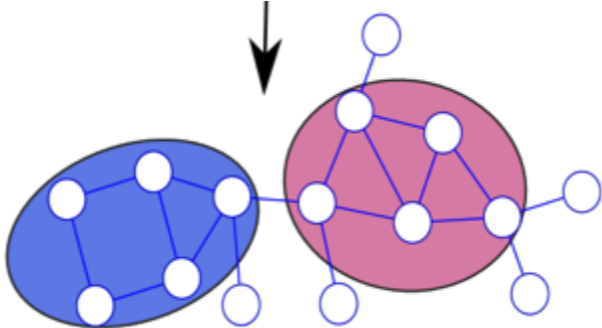
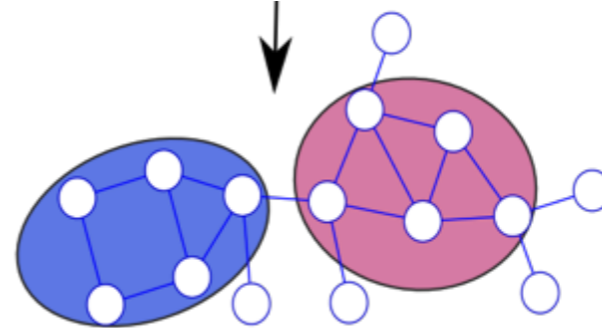
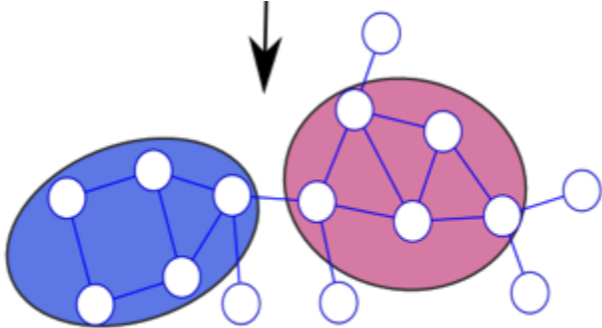
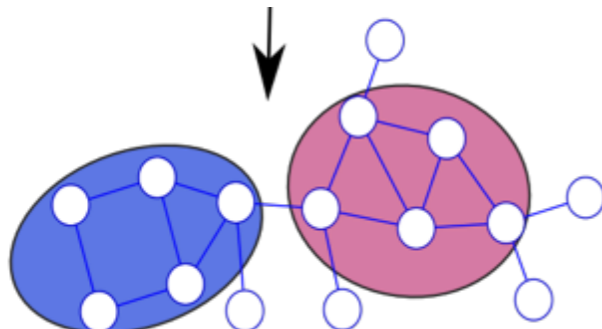
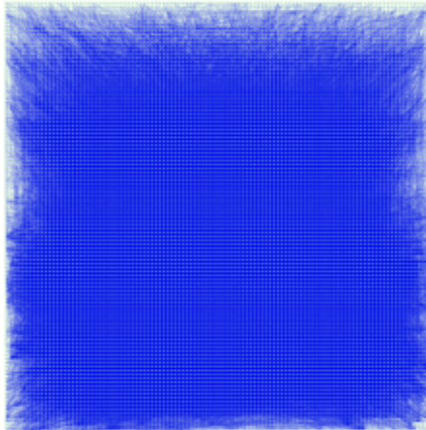
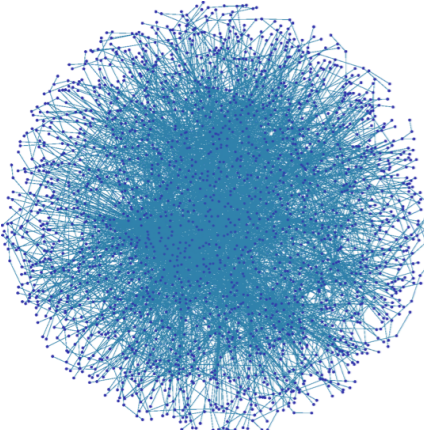
Union, Sum



Intersection



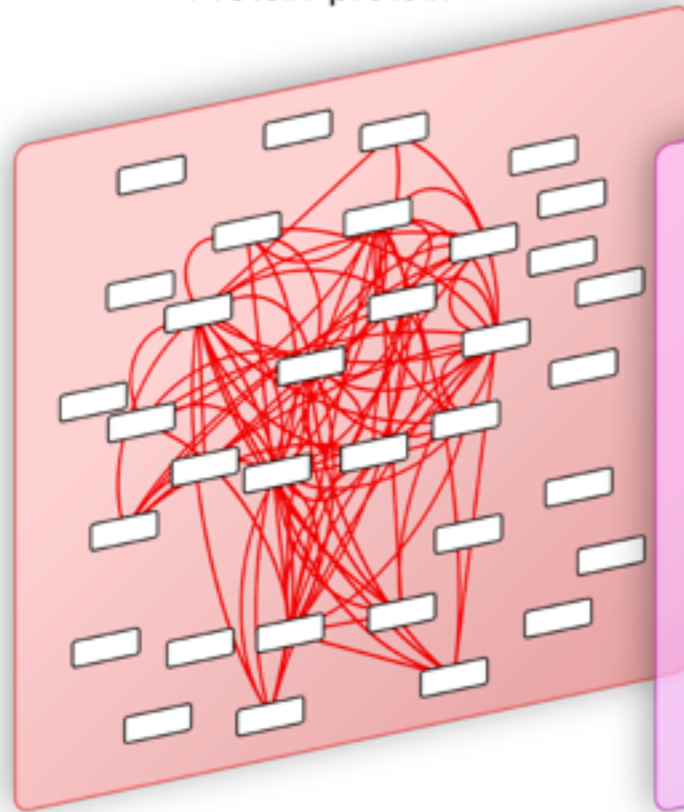
Consensus partitioning



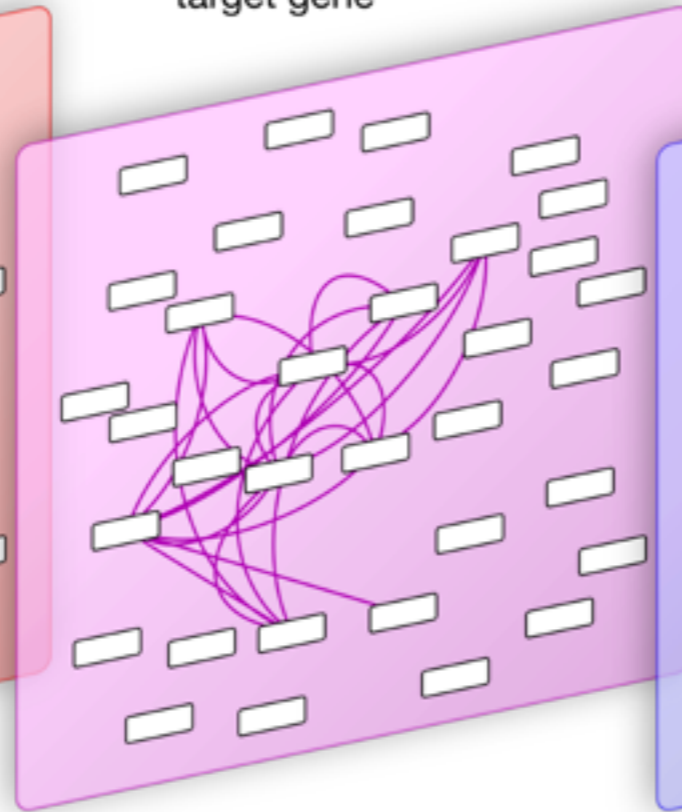


Multiplex framework

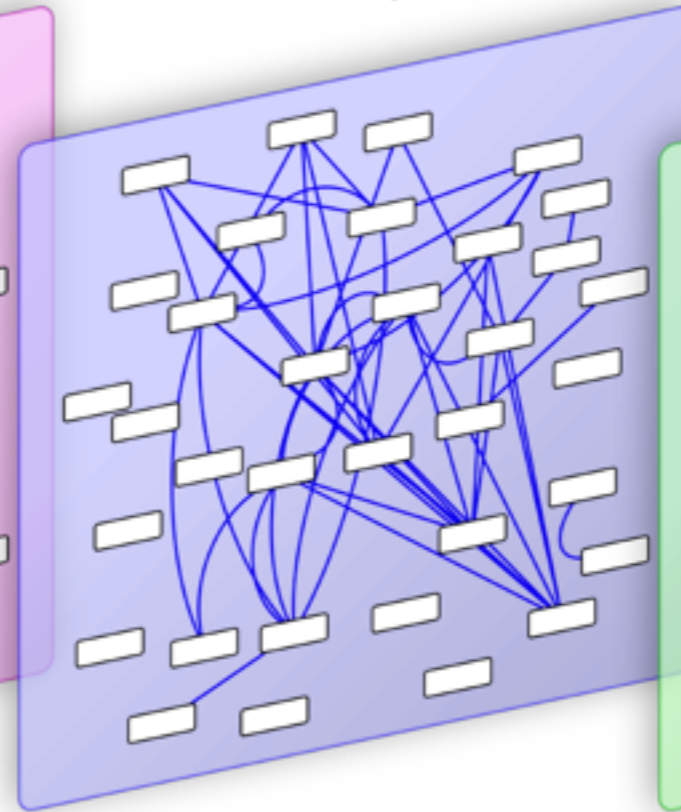
Protein-protein



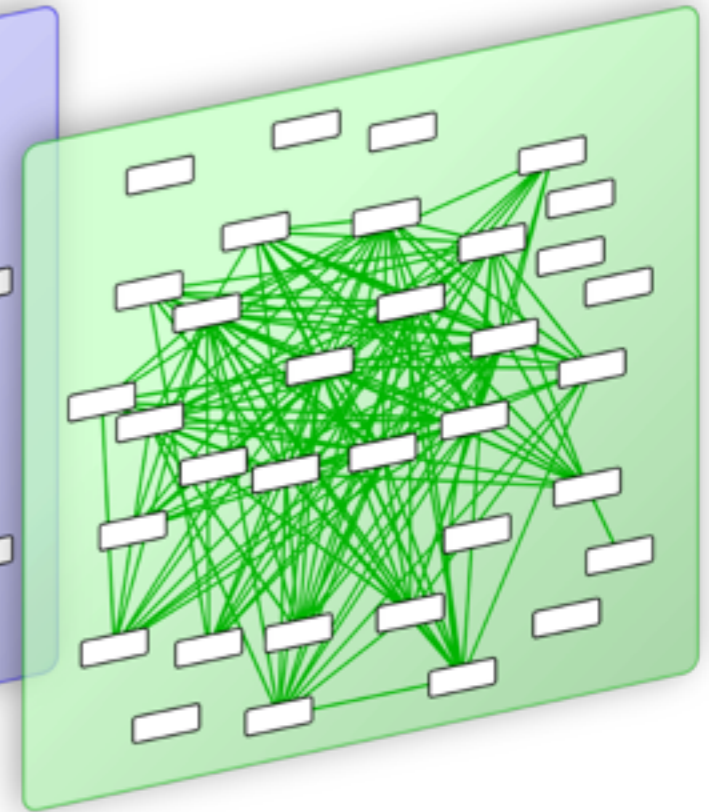
Transcription Factor - target gene



Pathways



Complexes

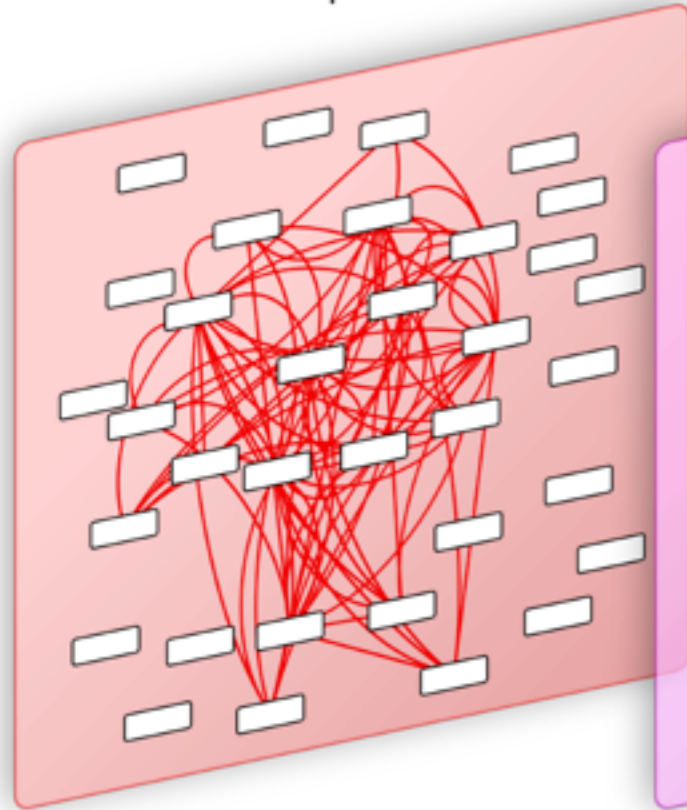


Multiplex, Multi-layer, Multislice

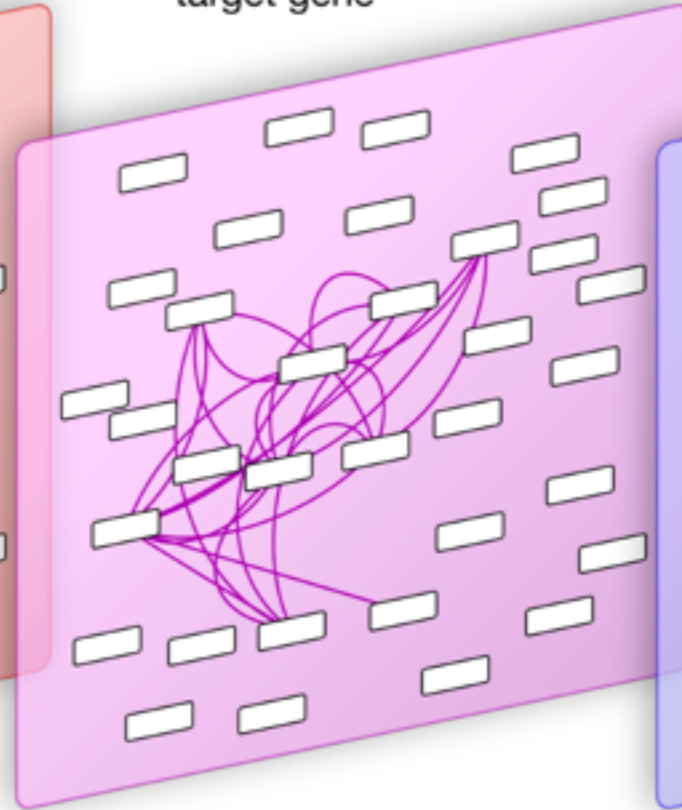


Multiplex framework

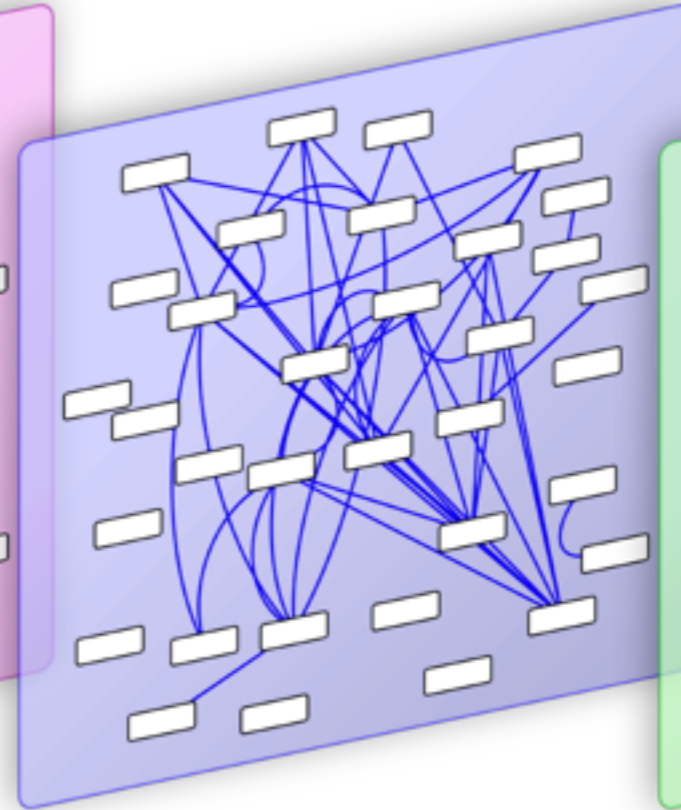
Protein-protein



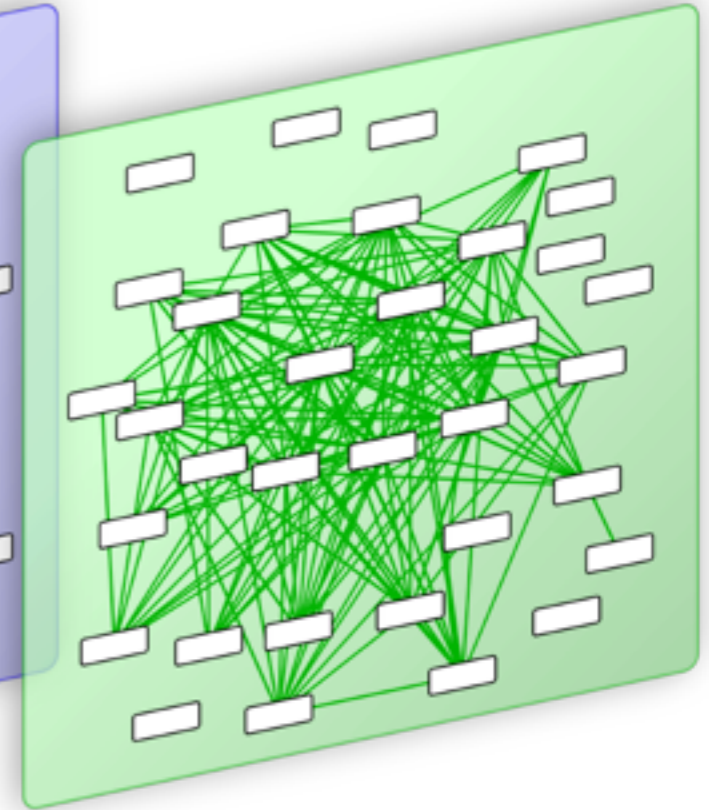
Transcription Factor - target gene



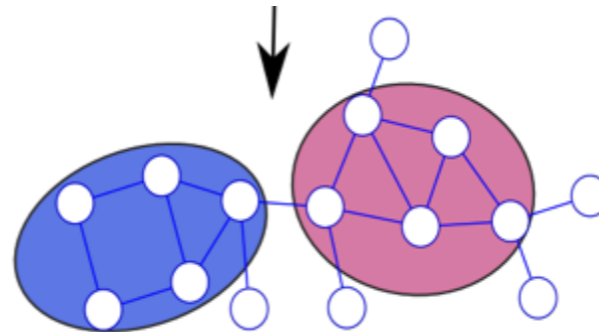
Pathways



Complexes



- **Multiplex-modularity**





Protocole

- **Multiplex framework**

- **Multiplex-modularity**
- **Mx-modularity Optimization (adapted Louvain algo)**

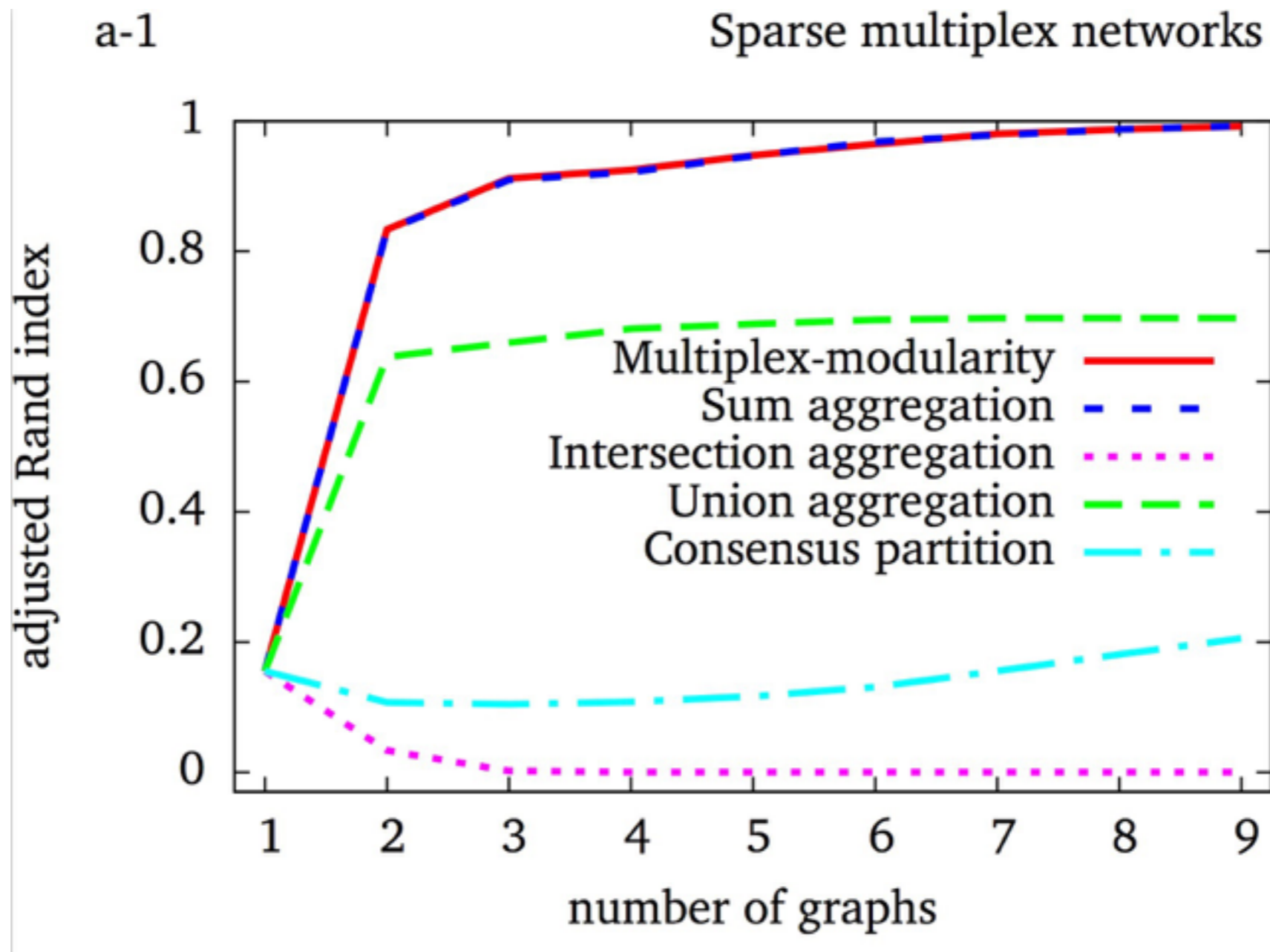
- **Networks aggregation**

- **Modularity**
- **Modularity Optimization (Louvain algo)**

- **Consensus partition**



Simulations (SBMs)



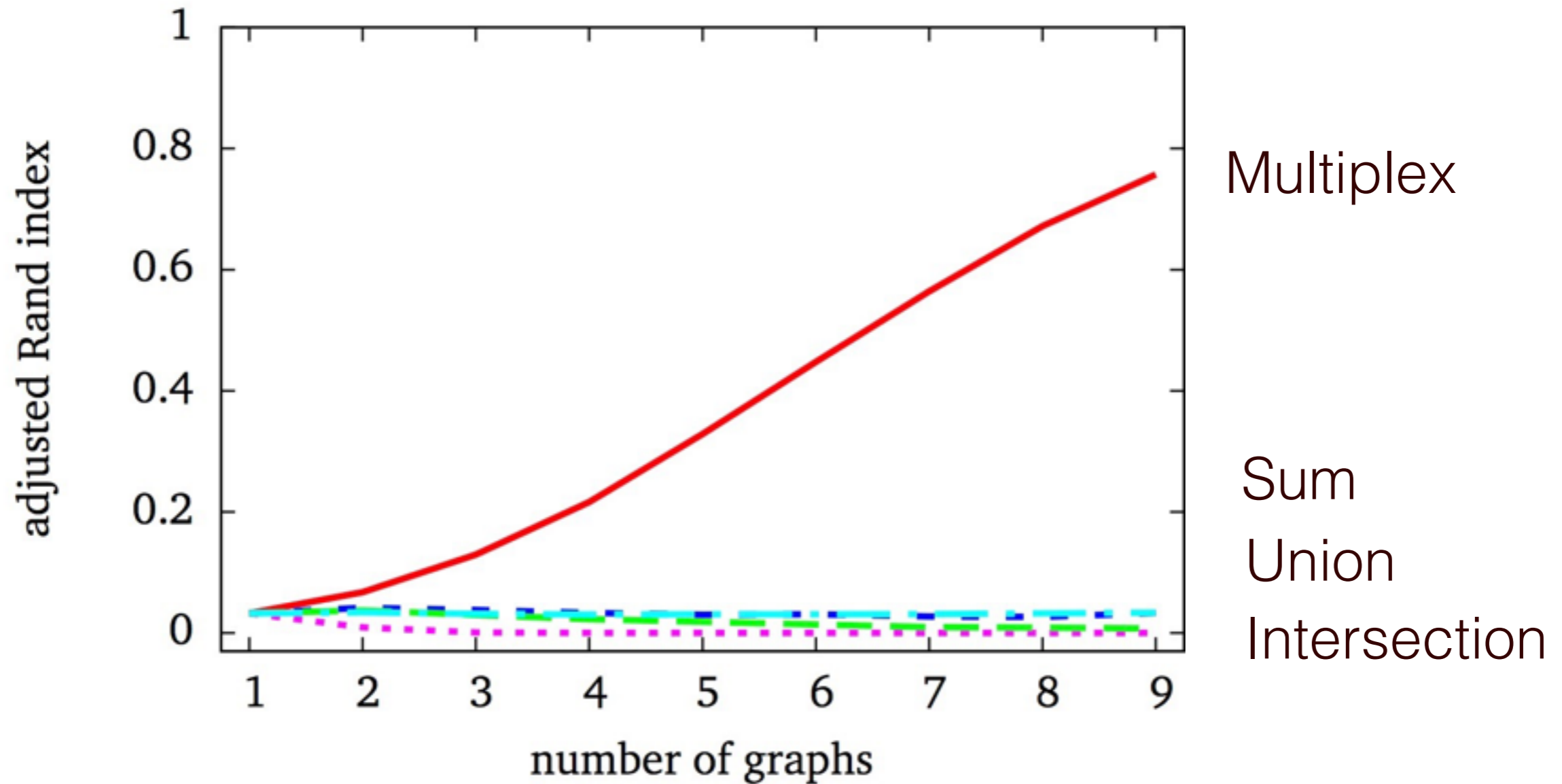
Considering many networks improves the detection of communities



Simulations (SBMs)

b-2

Mixed multiplex networks with missing data



Mx framework more efficient with heterogeneous and incomplete networks



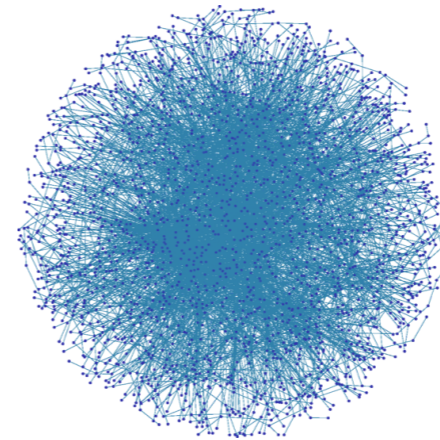
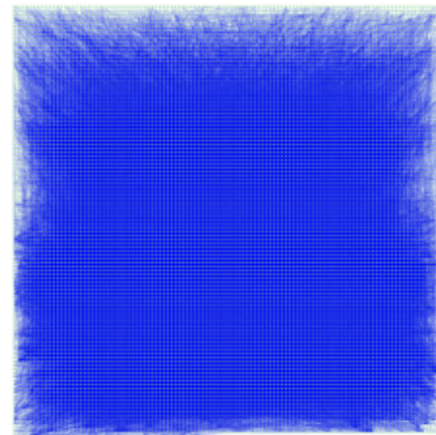
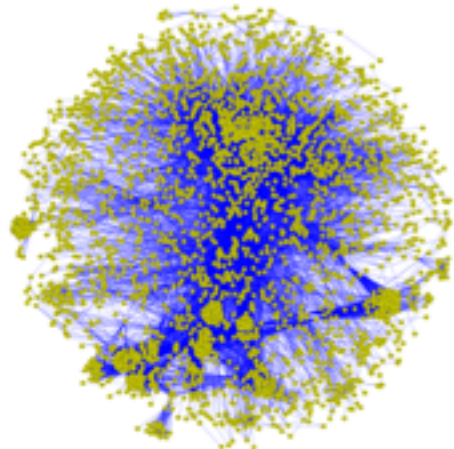
Application to real biological networks

Pathways

Co-expr

PPI

Complexes

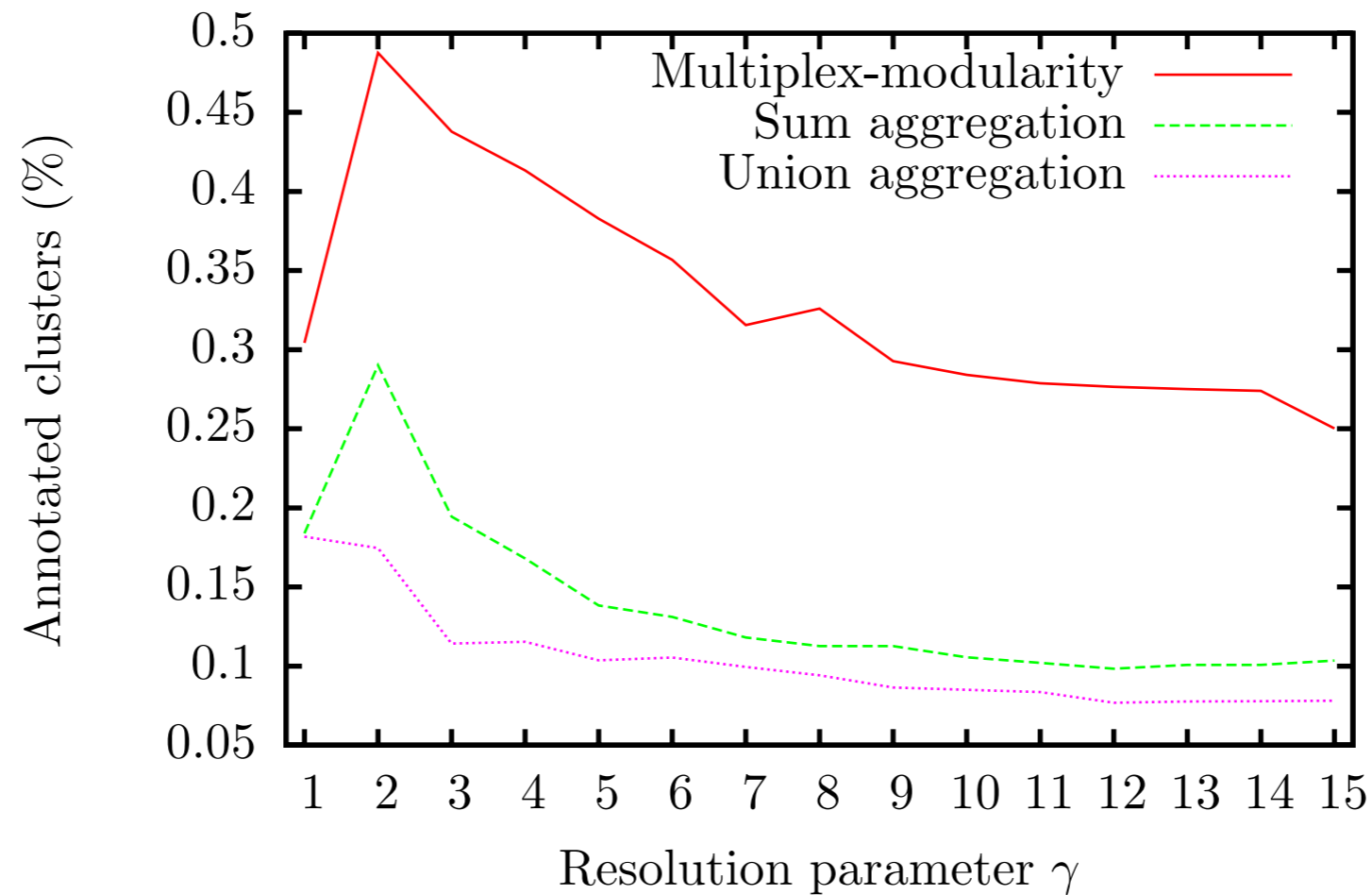


- Granularity parameter from 1 to 15
- Community number, sizes, similarities

=> More balanced



Community GOBP annotations



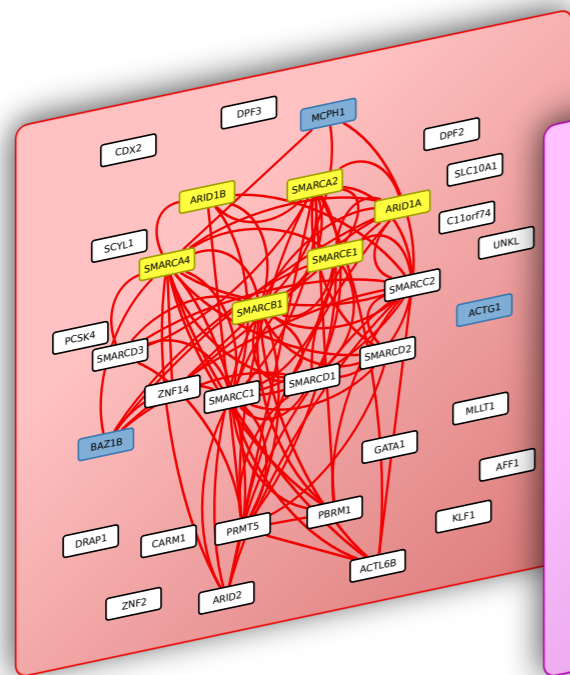
=> More annotated

=> Multiplex framework is well suited to identify communities from multiple biological networks

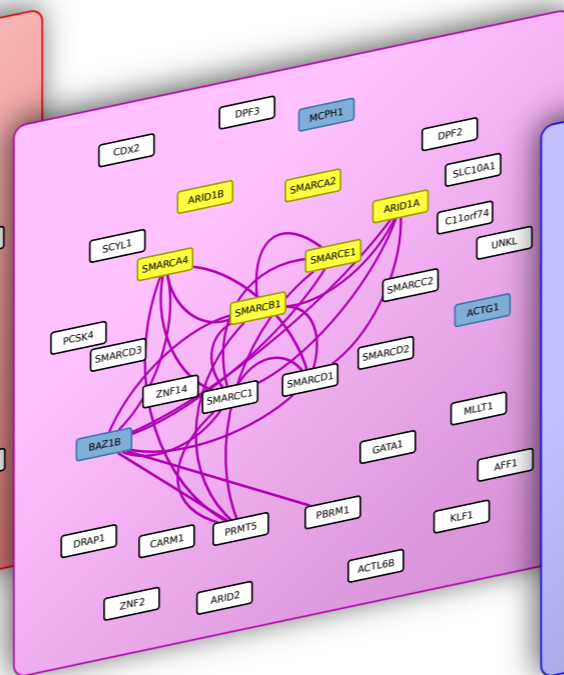


Module enriched in Coffin-Siris syndrome genes

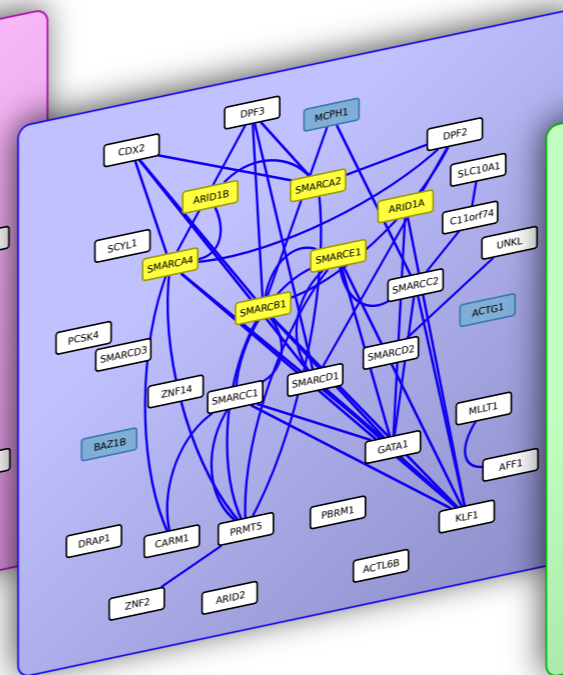
Pathways



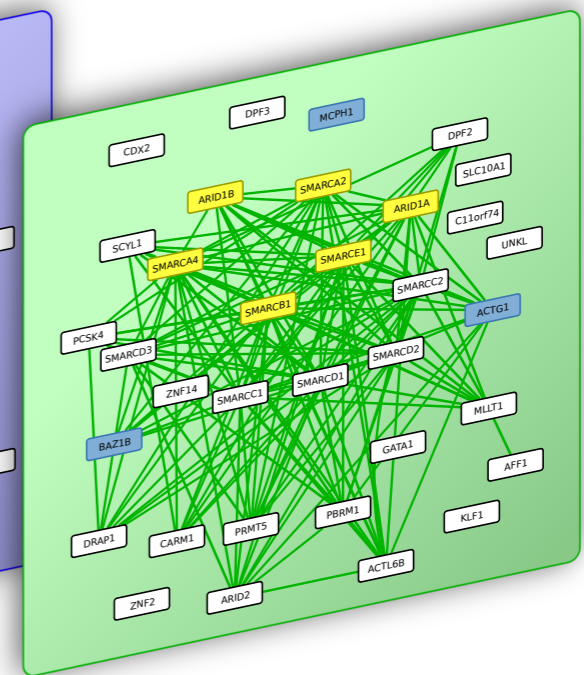
Co-expression



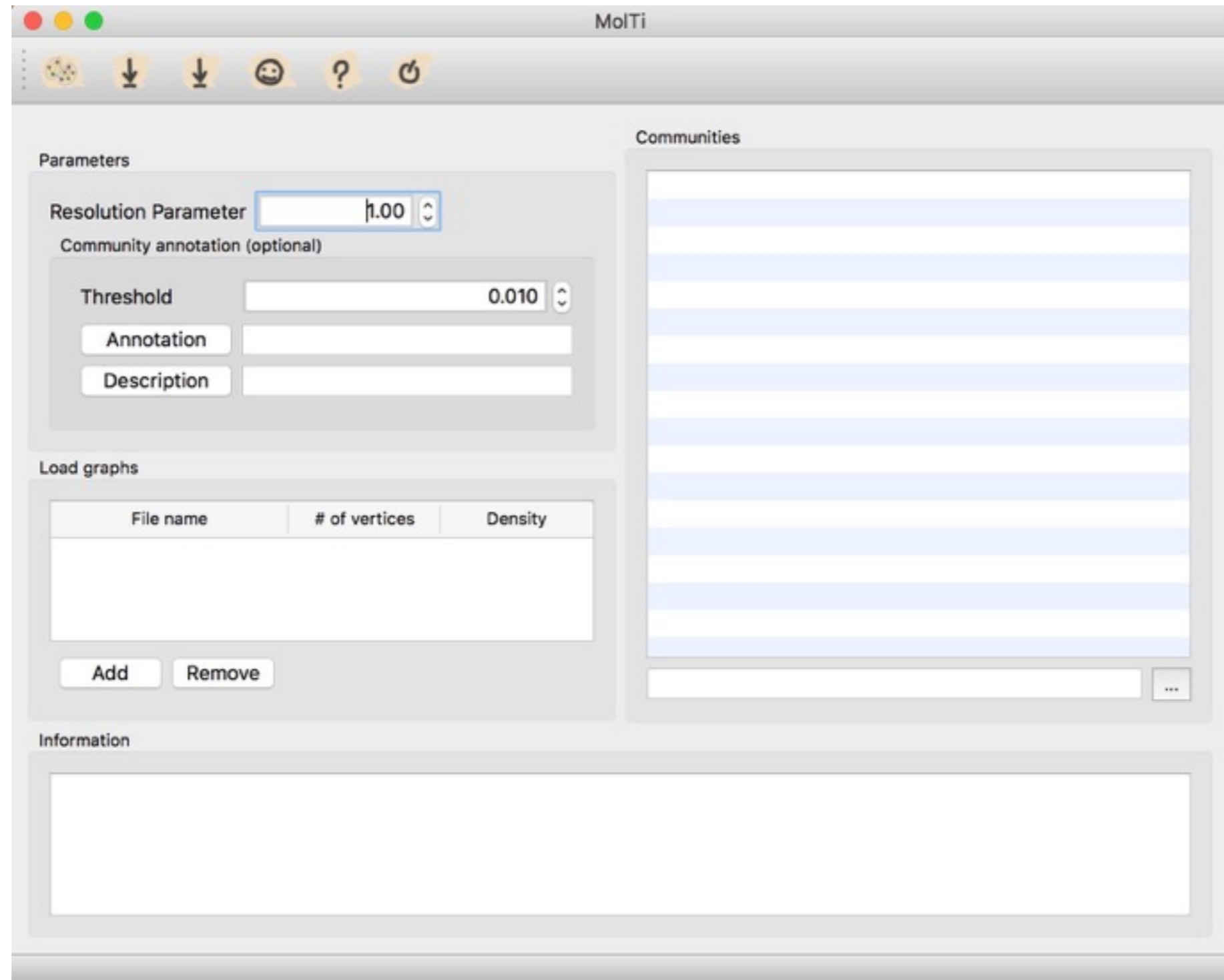
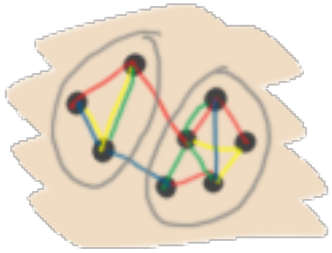
PPI



Complexes



Molti



<https://github.com/gilles-didier/Molti>



Acknowledgments



PEPS



Elisabeth Remy
Laurent Tichit
Alain Guénoche
Gilles Didier
José-Luis Portero