# ROTEIN-PROTEIN INTERACTION

DISEASE MECHANISMS TO THERAPEUTICS

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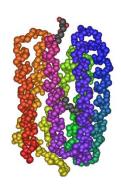




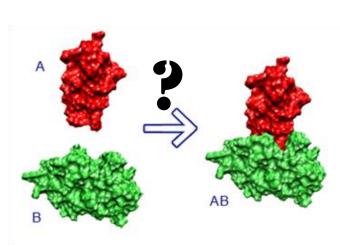


#### **Protein-Protein Interactions**

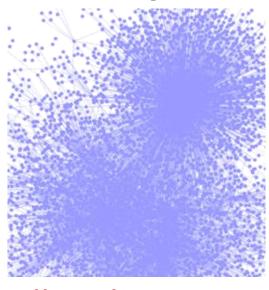
Central to all biological processes and structural scaffolds
The network is a medium for systems biology research
But 90% of protein-protein interactions are unknown today



~22,500 Protein-Coding Genes



Protein-Pairs ~250 Million



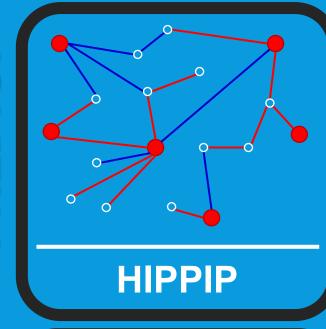
Human Interactome1 Million PPIs Estimated200k are known

### **Moonshots:**

Discover 100k novel PPIs and Achieve 100 Translational results <a href="Status"><u>Status</u>:</a>

Discovered 45k novel PPIs. 18 PPIs validated by experiment, all found to be true PPIs!

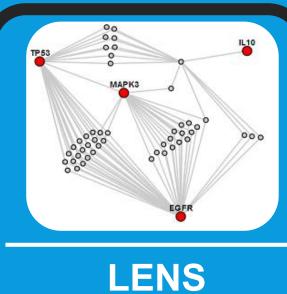
Achieved 2 high-impact translations.





Wiki-Pi

ANALY



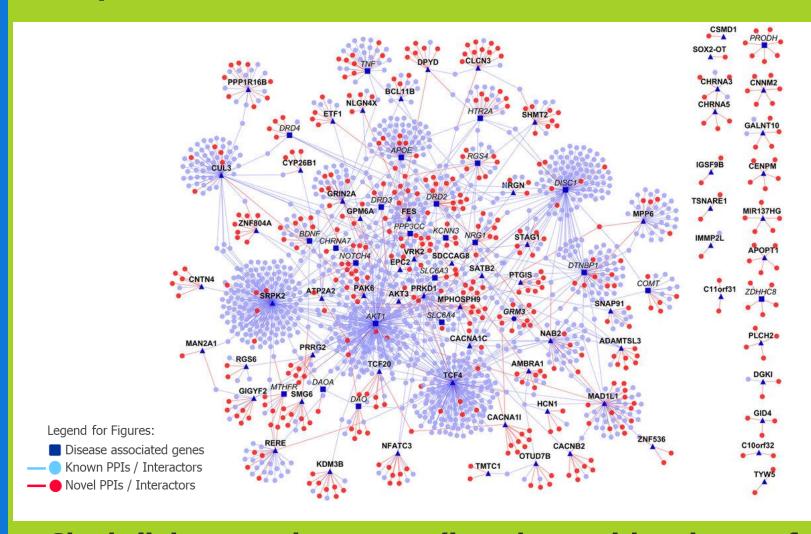


**BEST** 

PROTEIN-PROTEIN INTERACTIONS

## **Types of Contributions**

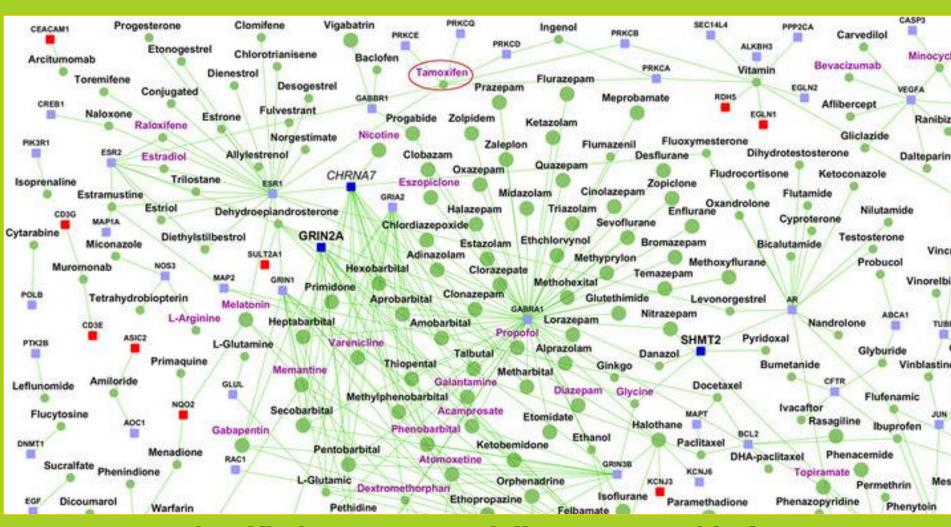
### **Study of Protein-Protein Interactomes of Disease-Associated Genes**



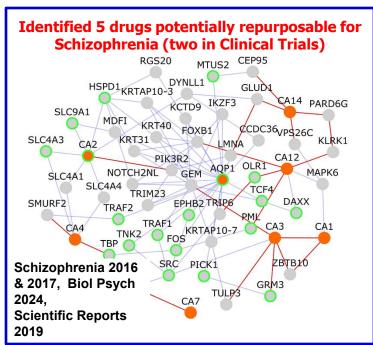
Sheds light on orphan genes (i.e., those with unknown functions)
Reveals unknown molecular connections to known pathways

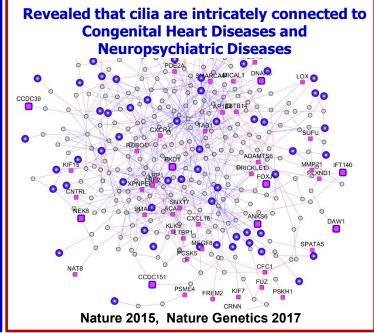
## **Types of Contributions**

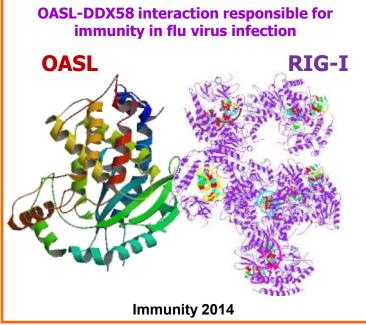
### Study of drugs targeting the network for repurposability



Identified Drugs Potentially Repurposable for Schizophrenia, and Mesothelioma. Two are in Clinical Trials for Scz.







Experimentally validated protein interactions. Every predicted PPI that was studied experimentally was found to be valid.

1	STT3A	RPS25	6	ALB	KDR	11	CLPS	CUTA
2	STT3A	SYCP3	7	ALB	PDGFRA	12	HMGB1	CUTA
3	STT3A	MCAM	8	STX3	LPXN	13	HMGB1	FLT1
4	DDX58	OASL	9	STX4	MAPK3	14	HMGB1	KL
5	BAP1	PARP3	10	WDR5	IGFBP3	15	IFT88	KL

https://www.nature.com/articles/npjschz201612

https://www.nature.com/articles/nature14269

https://www.nature.com/articles/ng.3870

https://www.nature.com/articles/s41537-017-0010-z

https://www.nature.com/articles/s41598-019-48307-w

https://www.sciencedirect.com/science/article/pii/S1074761314001873

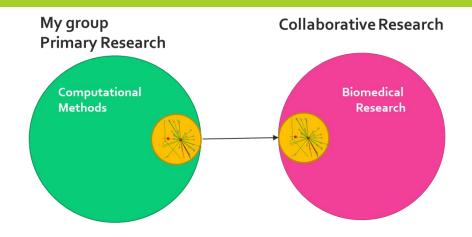
Clinical Trials: https://clinicaltrials.gov/ct2/show/NCT03794076

(2<sup>nd</sup> with IRB review)

**Computationally Predicted PPIs are making Translational Impact** 

## **Types of Contributions**

### **Translation to Bench and Bedside through Collaboration**





PPIs are studied in labs, and drugs are being tested in clinic.