

Analyzing genetic risk locations from GWAS in relation to vital genes and pathways using QIAGEN Ingenuity Pathway Analysis



Grace Chou 周儀柔 資深業務專員

Bryce Lin 林文賢 經理

[Bioinfo@GGA.ASIA](mailto:Bioinfo@GGA.ASIA)

- Introduction to QIAGEN Ingenuity Pathway Analysis
- Analyzing risk loci using IPA
  - Data format
  - Data upload and analysis setup
  - Core analysis
- Comparing between risk loci analysis
  - Comparison Analysis
  - Compare
- Querying IPA's Knowledge base
  - Custom network construction for risk loci associated genes
  - Contextualization using public data
- Summary

Account: ipaXX@ingenuity.com

(XX= 01~45)

Password: **9-Str0ngP@&&IPA**

Expired until 10/20

*Download website:* <https://tinyurl.com/GGAIPA>

*Dataset:* <https://tinyurl.com/GGAdataset>

*Client: (IPA Login)*



# IPA Download webpage



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## Ingenuity Pathway Analysis

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## Choose your IPA installer

Ingenuity  
Pathway  
Analysis



### Install IPA on your computer

This installer will enable you to access IPA like other desktop applications on your computer (though still requiring an internet connection).

Note: This is all you need to run IPA. It is not necessary to install Java separately from IPA.

Click on the button below to download the installer

**IPA for Windows (64-bit)**

(Installer recommended for your computer)

Other options:

[IPA for 32-bit Windows](#)

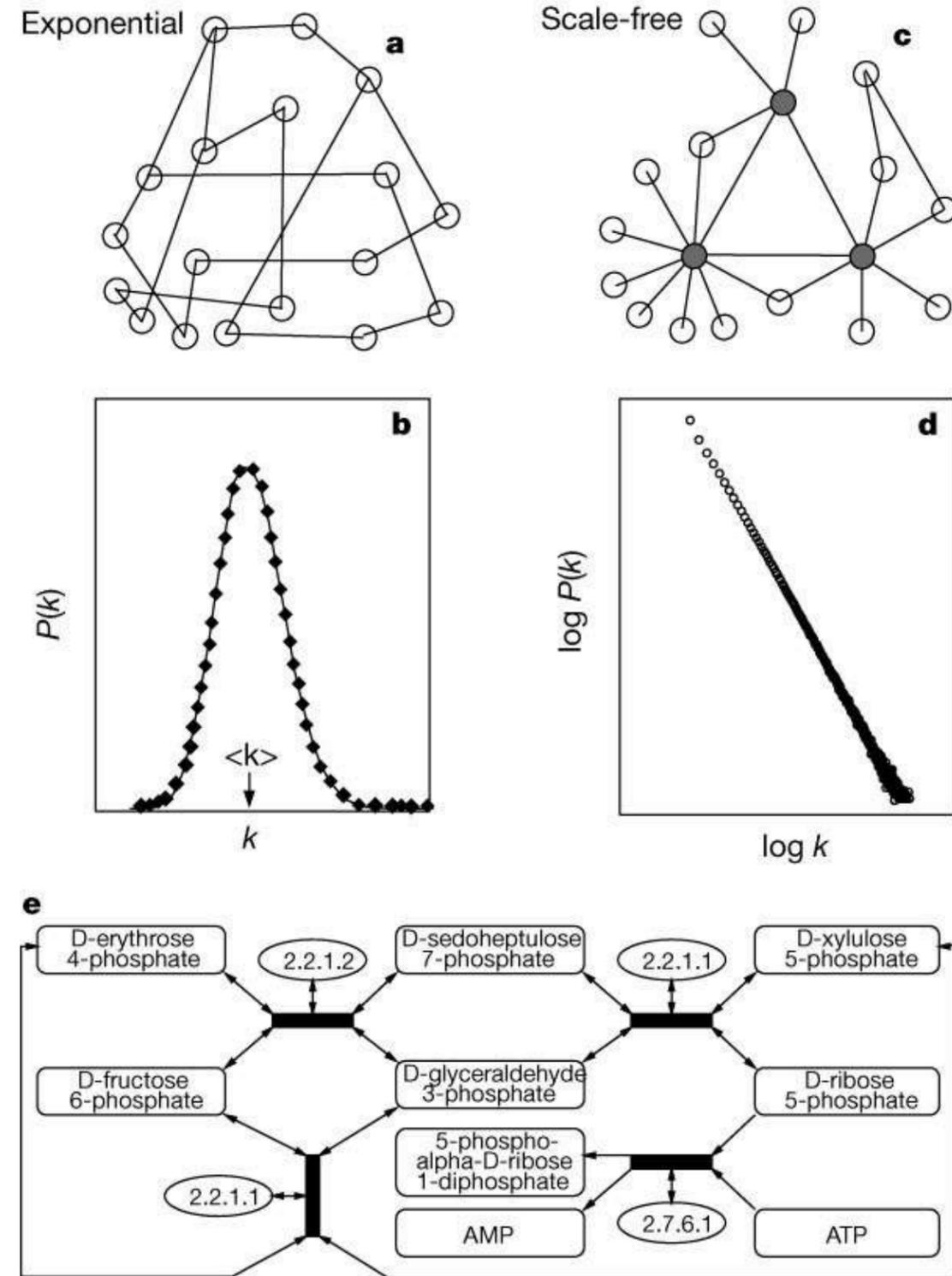
[IPA for macOS](#)

For more information see this [help page](#).

Note: If you have trouble installing or logging in with the IPA client, please click [here](#) for an alternate method to launch IPA.

Looking for more information about IPA? Click [here](#).

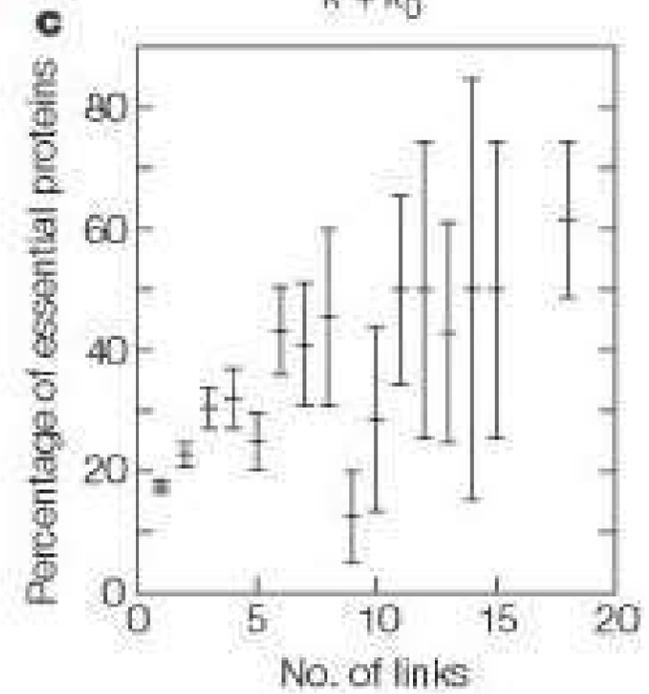
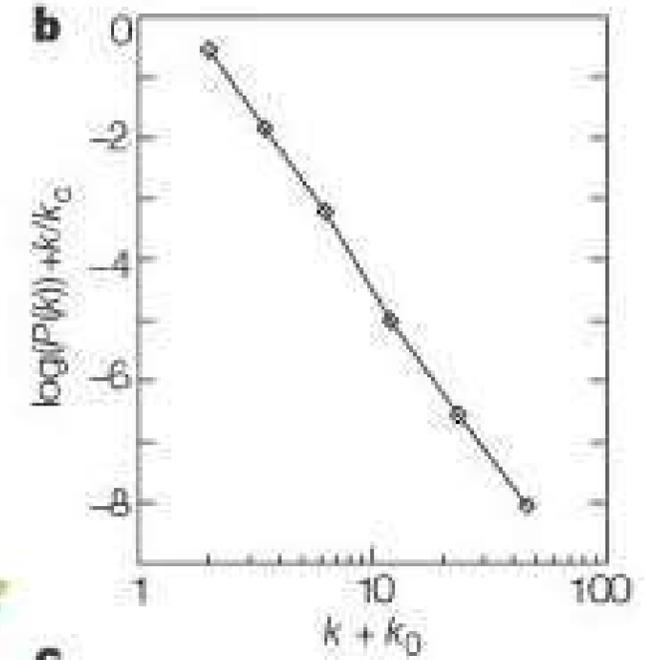
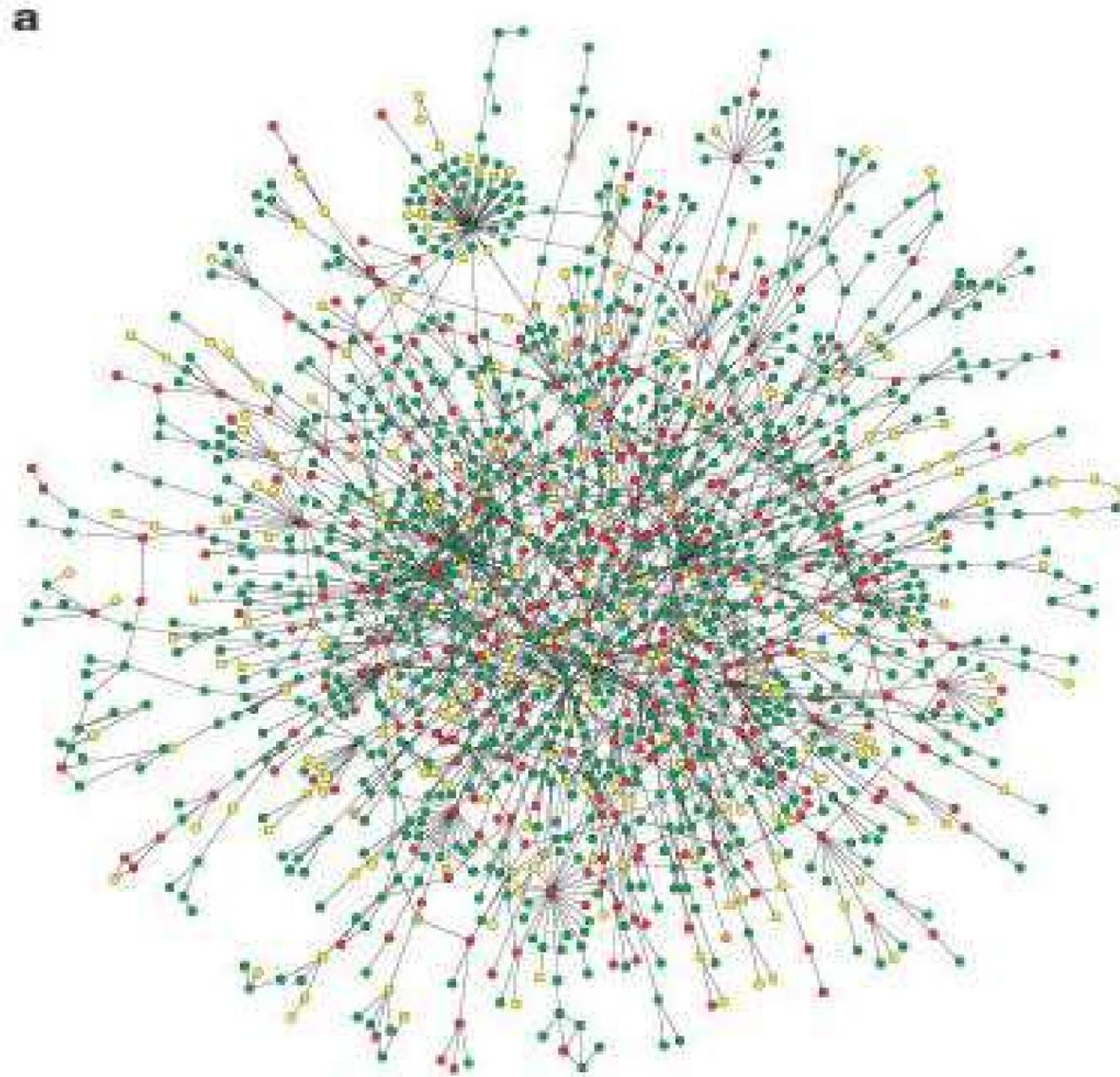
Despite significant variation in their individual constituents and pathways, these metabolic networks have the same topological scaling properties and show striking similarities to the inherent organization of complex non-biological systems. This may indicate that metabolic organization is not only identical for all living organisms, but also complies with the design principles of robust and error-tolerant scale-free networks, and may represent a common blueprint for the large-scale organization of interactions among all cellular constituents.



[Nature 407, 651-654 \(2000\)](#)

The most highly connected proteins in the cell are the most important for its survival.

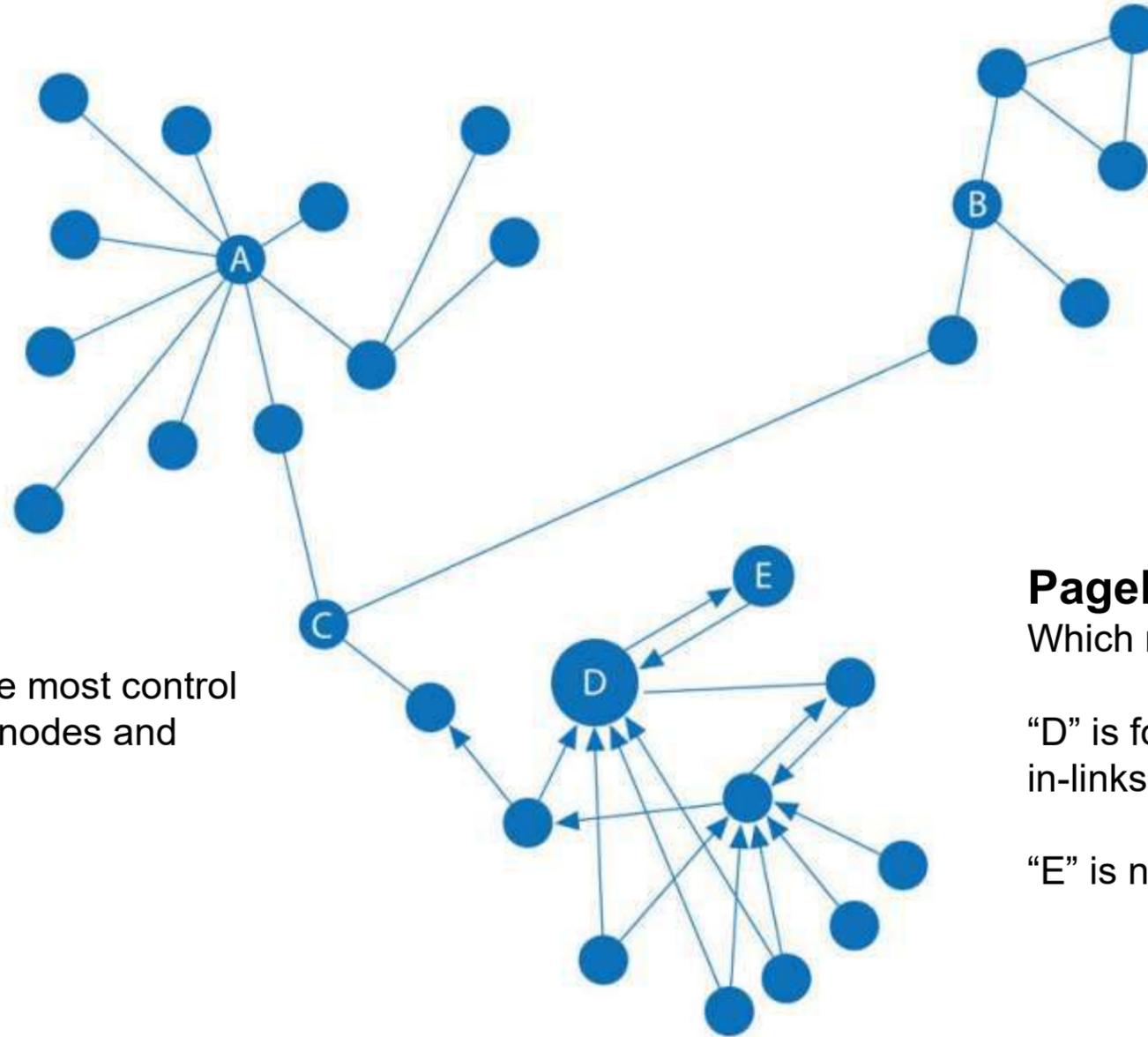
[Nature 411, 41-42 \(2001\)](#)



## Degree

Number of connections?

“A” has a high degree



## Closeness

Which node can most easily reach all other nodes in a graph or subgraph?

“B” is closest with the fewest hops in its subgraph

## Betweenness

Which node has the most control over flow between nodes and groups?

“C” is a bridge

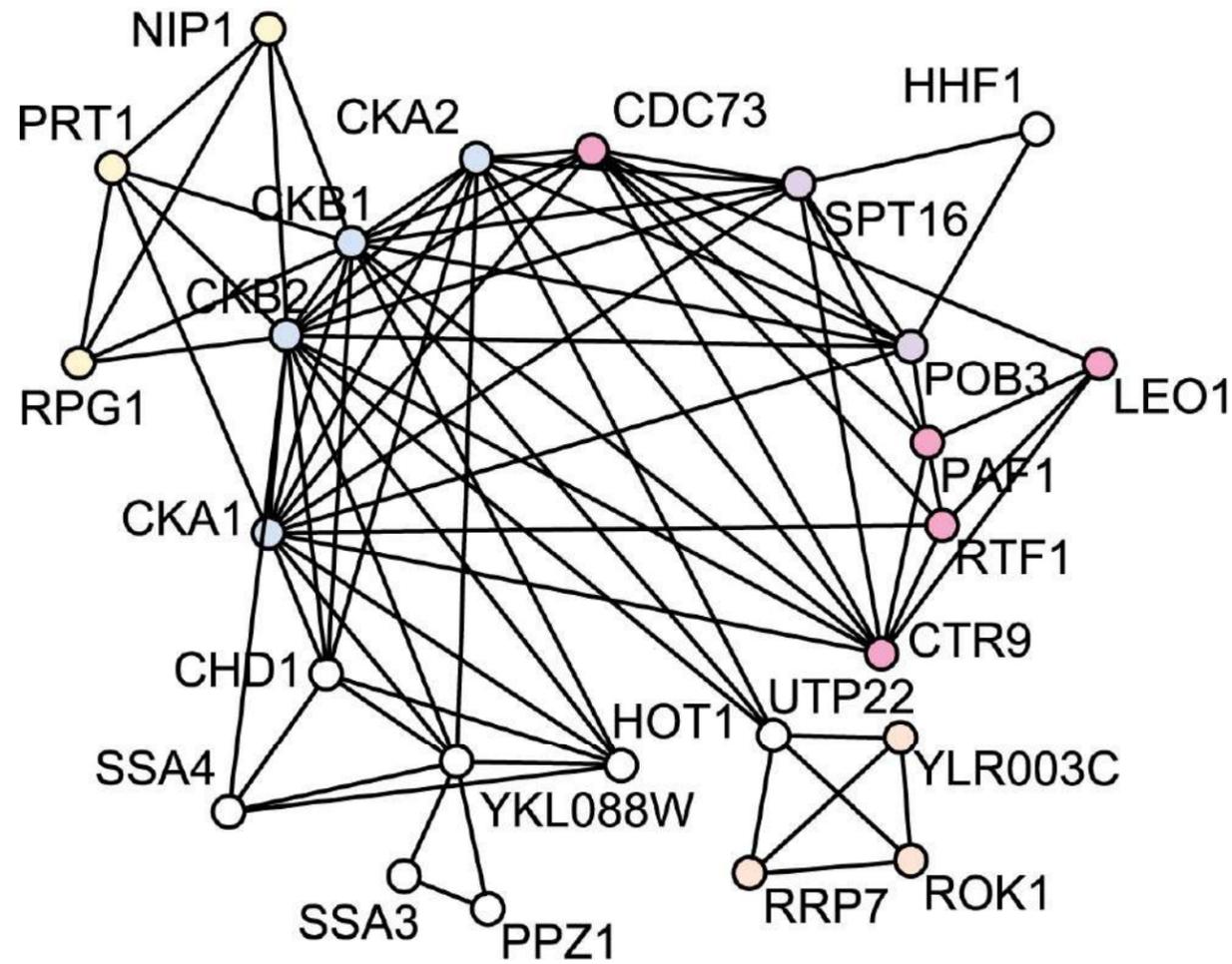
## PageRank

Which node is the most important?

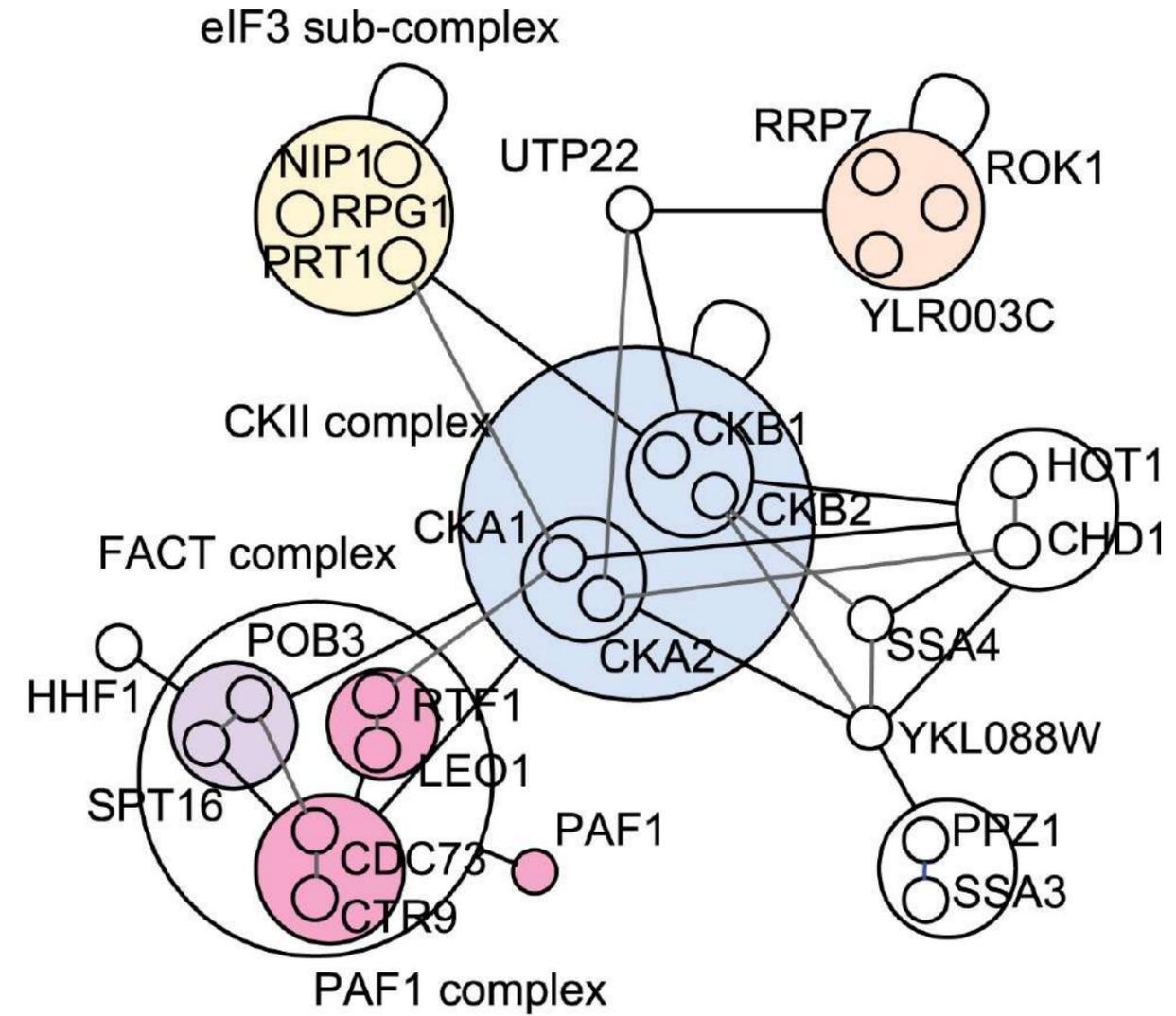
“D” is foremost based on number & weighting of in-links

“E” is next, due to the influence of D’s link

(Source: [Needham & Hodler, 2019](#))

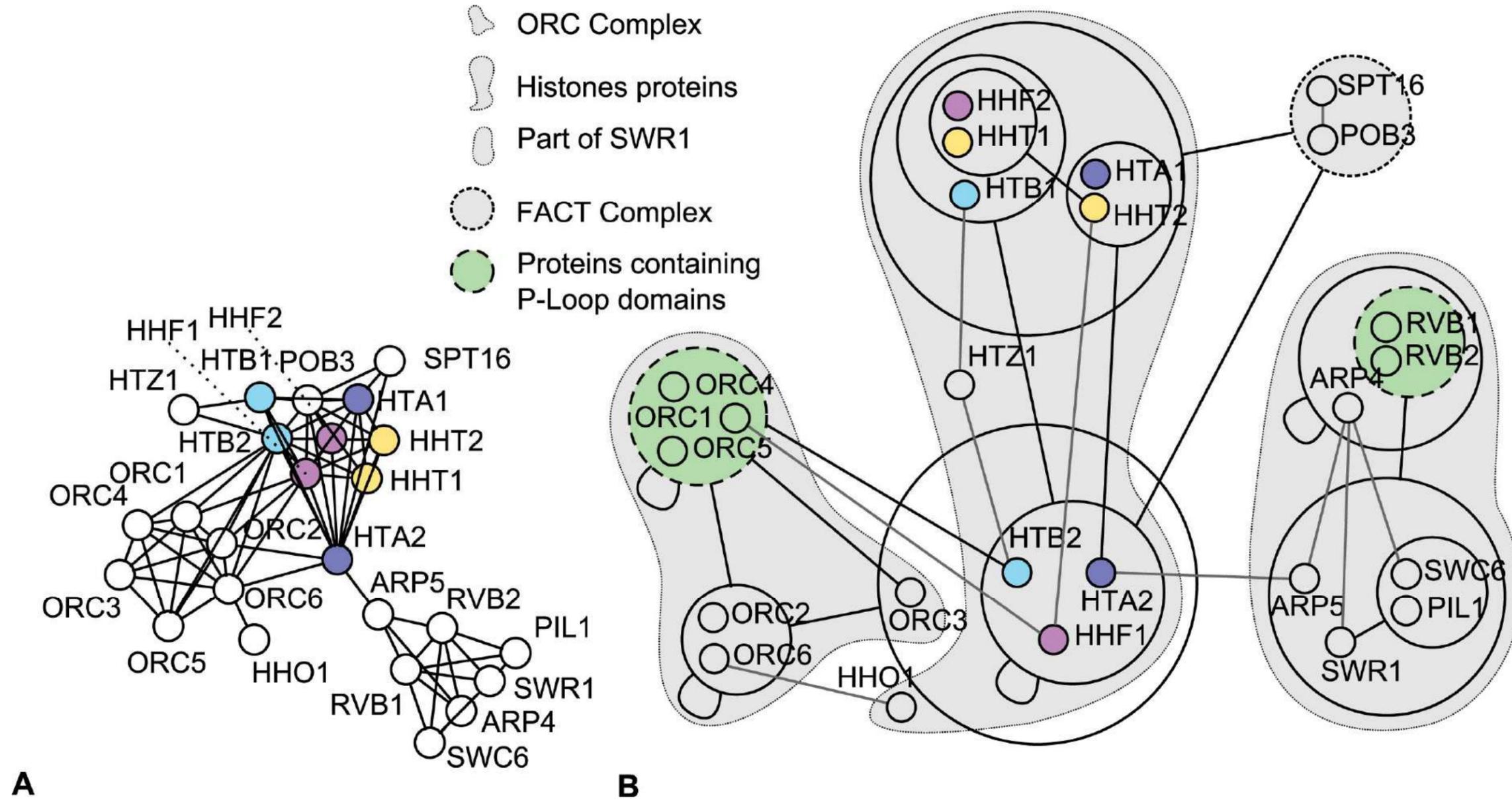


**A**



**B**

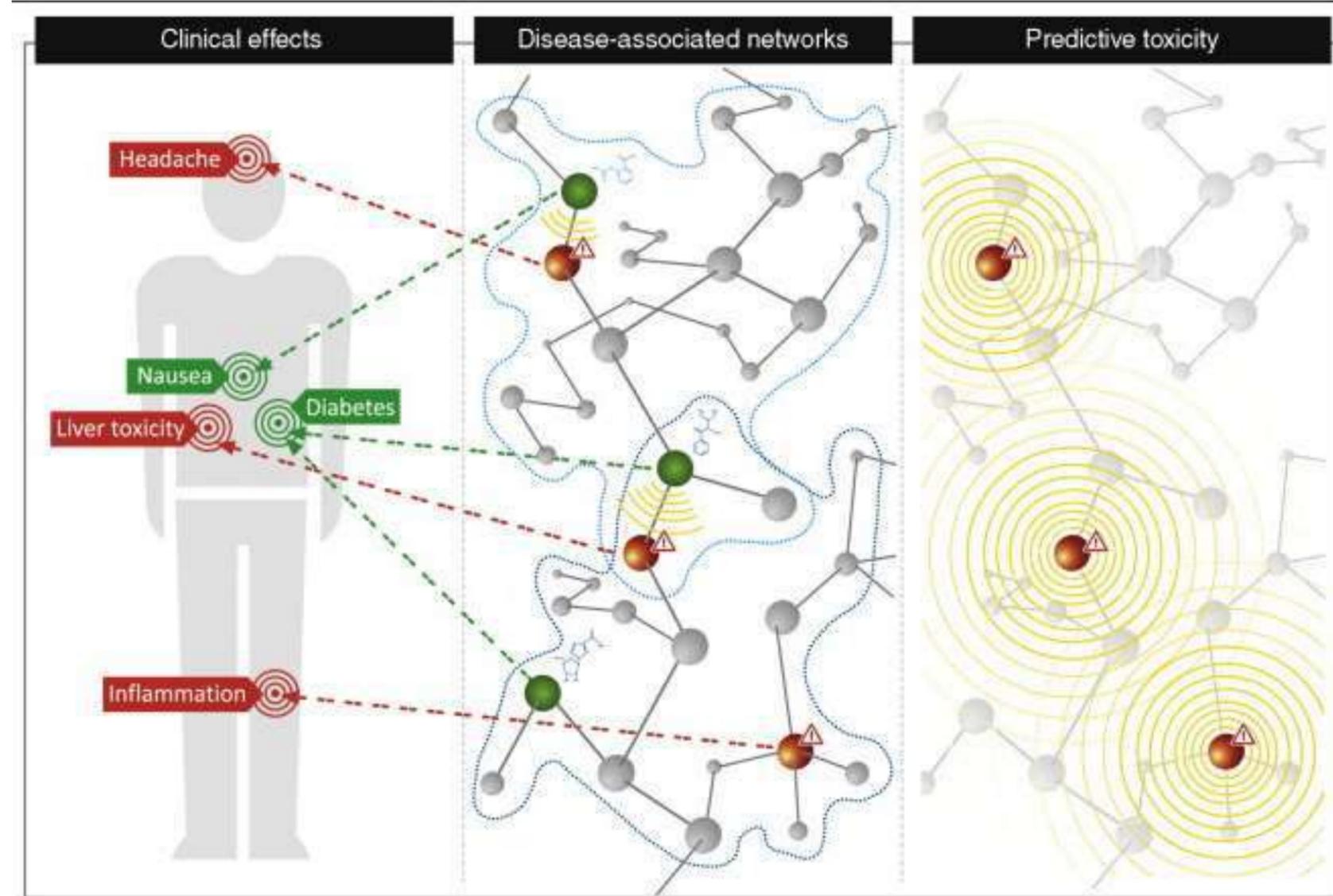
•<https://doi.org/10.1371/journal.pcbi.1000108>



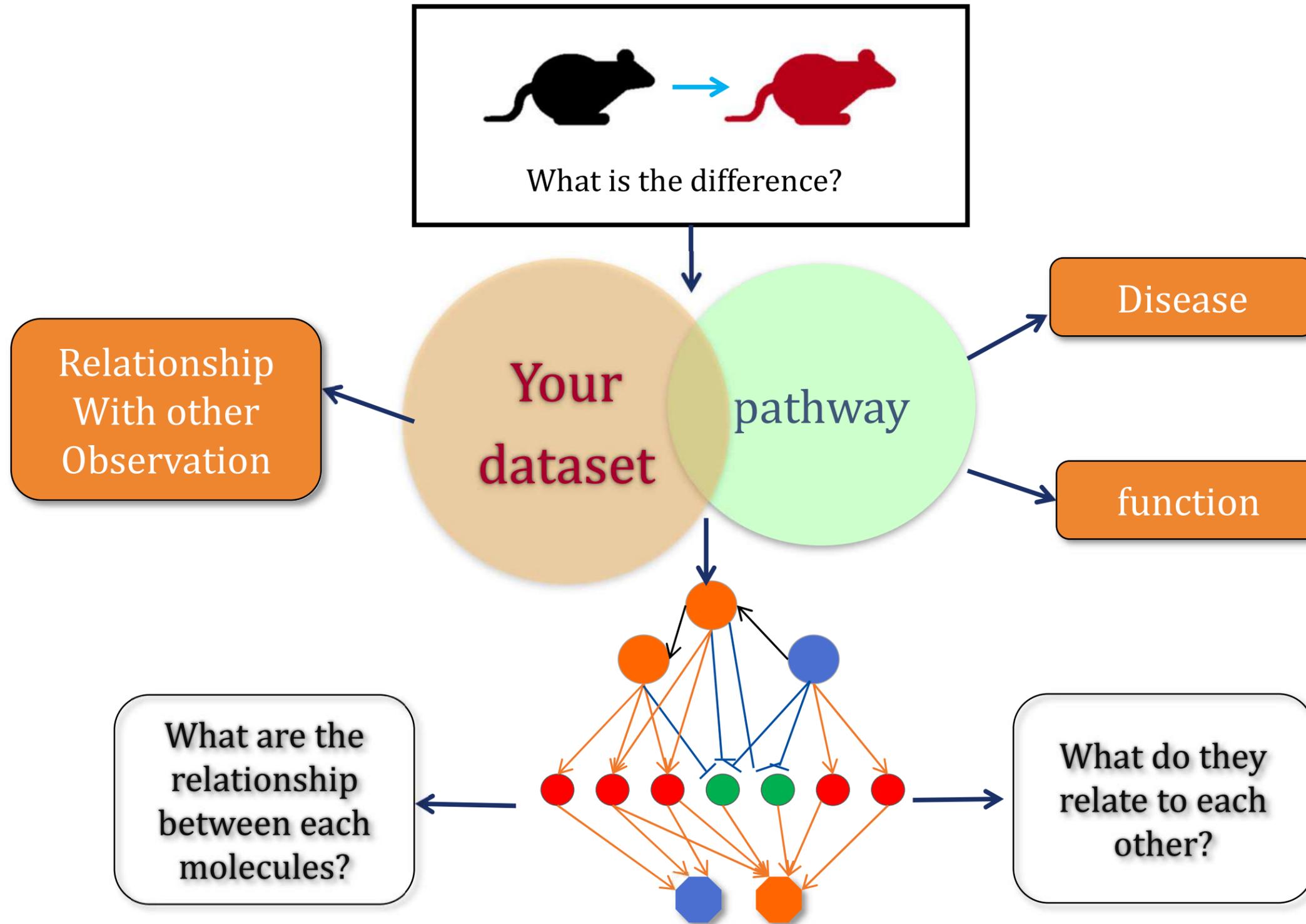
•<https://doi.org/10.1371/journal.pcbi.1000108>

Models can identify the areas of influence of proteins leading to undesired effects, and to explore how they are related to network connectivity.

- Identifying potential drug targets
- Designing the toxicity tests
- Highlighting the existence of validated drug targets for a given therapeutic indication in key enclaves of the network describing a different disease, thereby suggesting candidates for drug repurposing (i.e. finding new indications for a target).



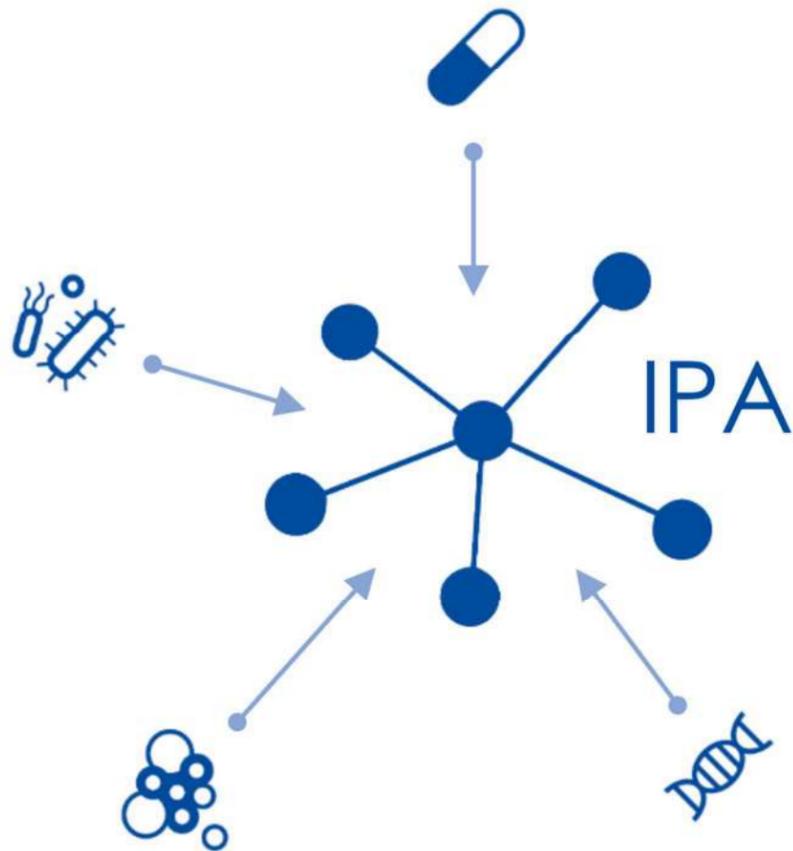
TRENDS in Pharmacological Sciences



Updated weekly with the latest clinical trials and manuscripts

12+ million findings curated using controlled vocabulary

Decades of accumulated knowledge mapped in directional networks

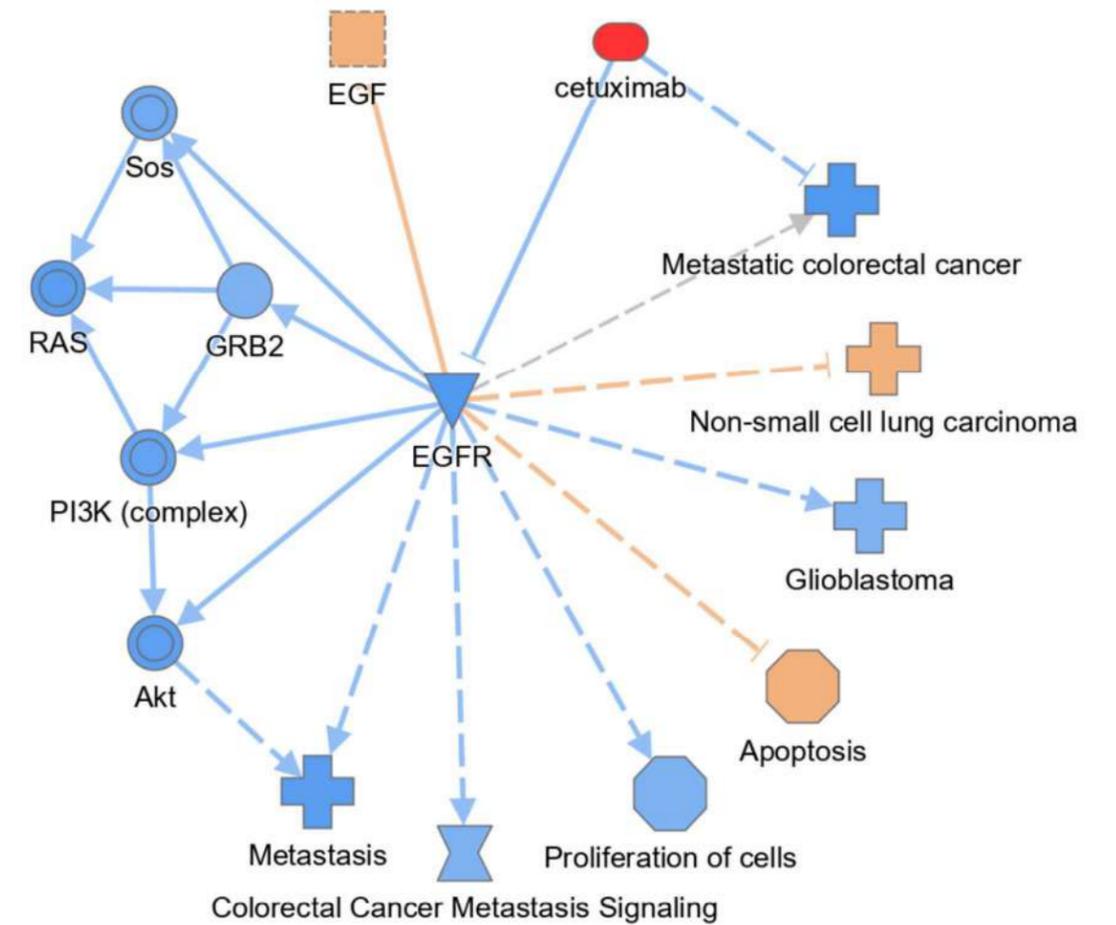


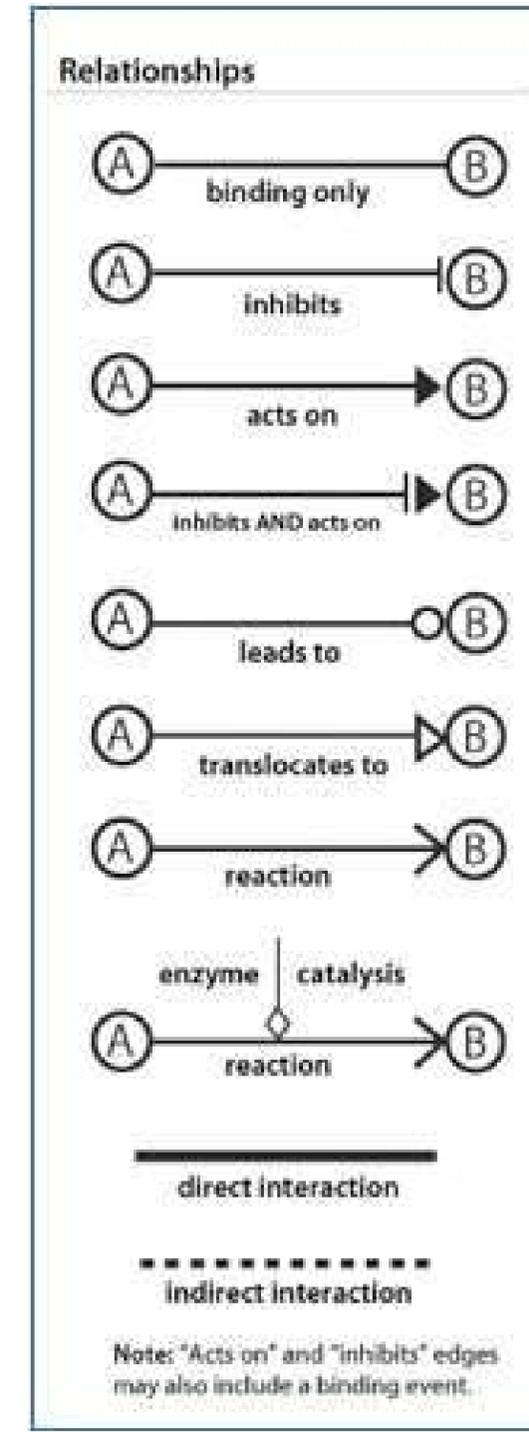
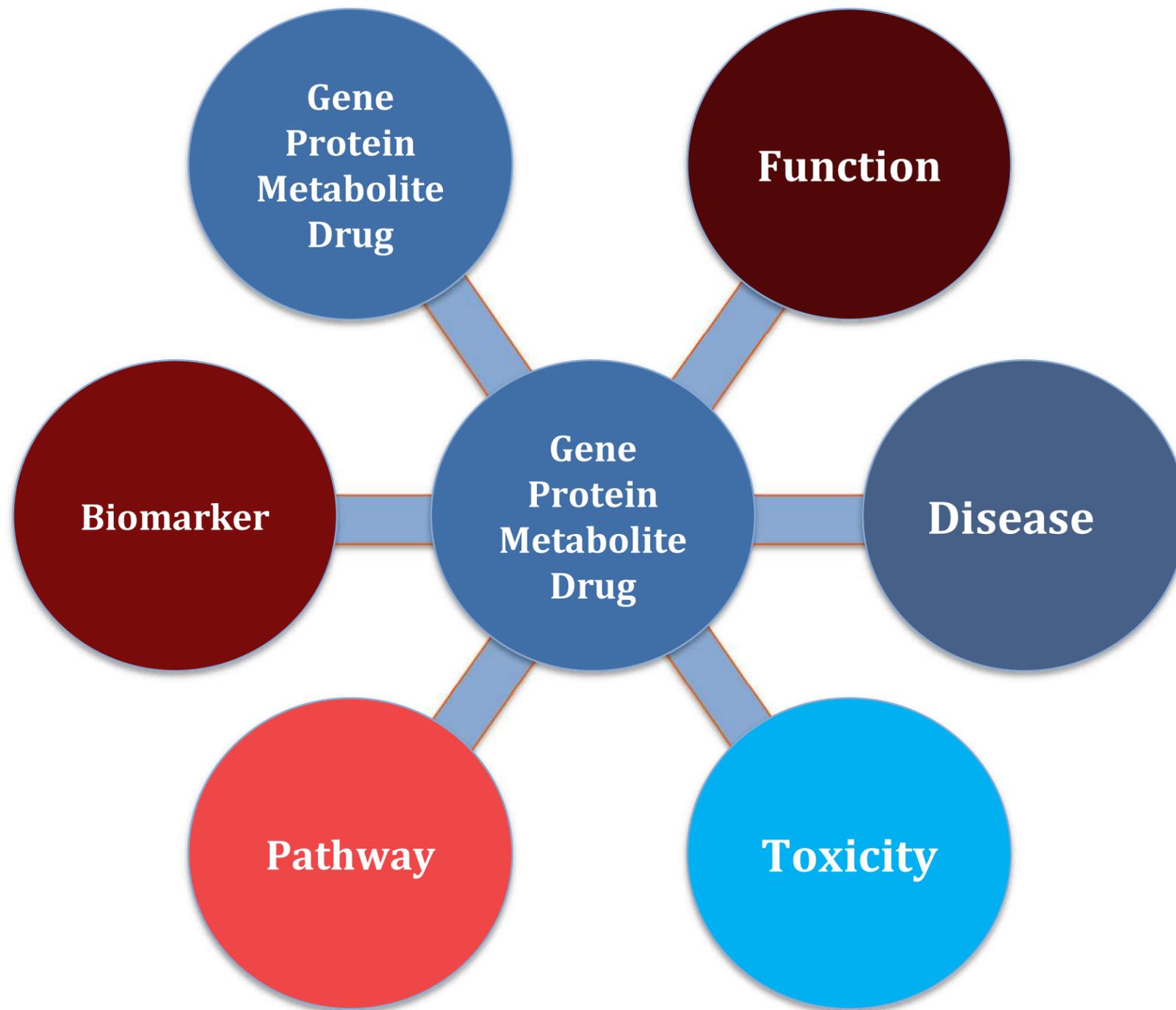
**EGF – EGFR**  
[one of many]

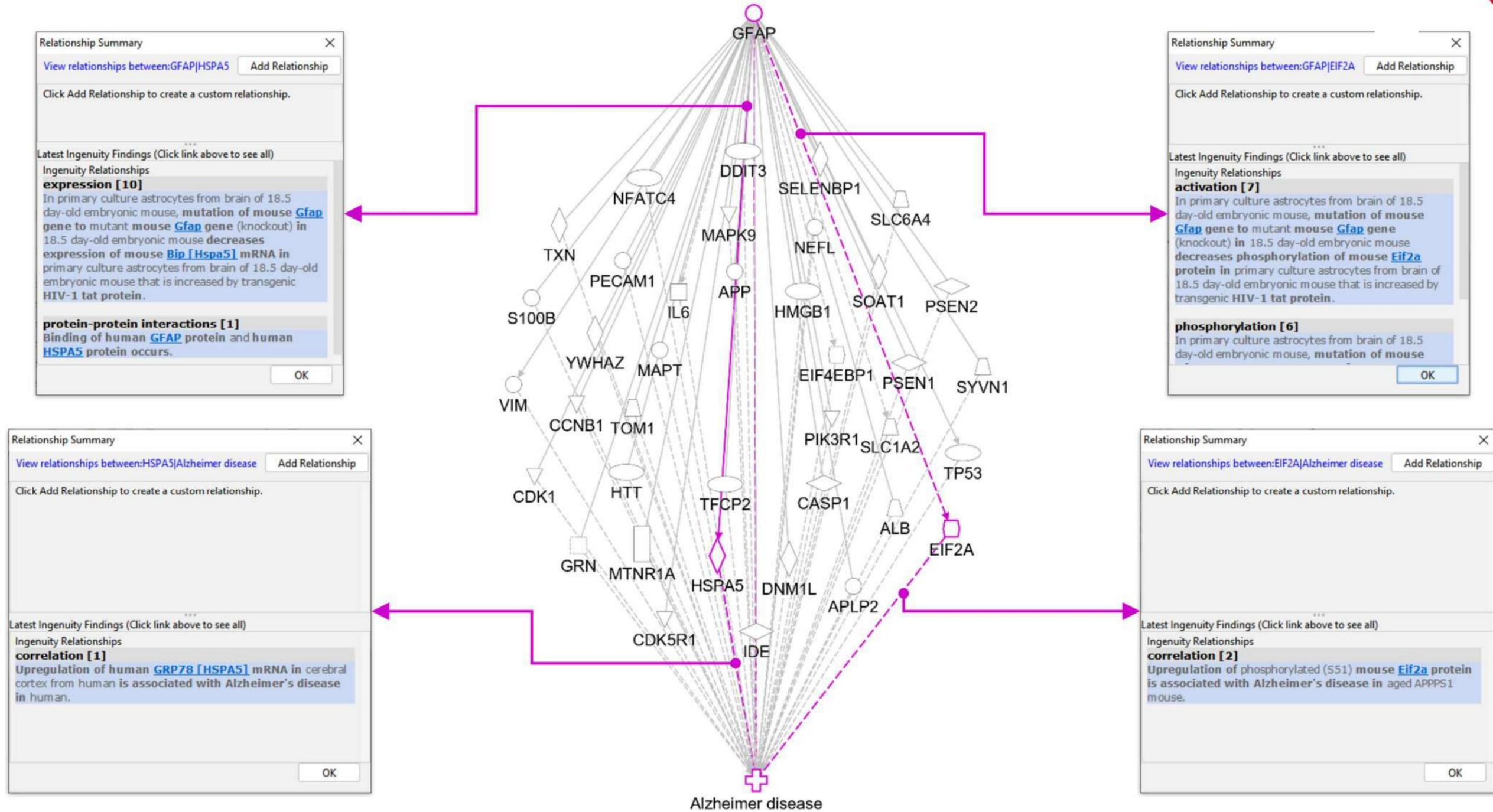
Type: **activation**  
 Direction: **directional**  
 Effect: **increases**  
 Directness: **direct**  
 Tissue: **epithelial cells**  
 Location: **plasma membrane**  
 Source: **PMID: 17909010**

**cetuximab – EGFR**  
[one of many]

Type: **phosphorylation**  
 Direction: **directional**  
 Effect: **decreases**  
 Cell line: **CaR1 cells**  
 Organism: **human**  
 Experiment: **immunoblot**  
 Source: **PMID: 23213241**

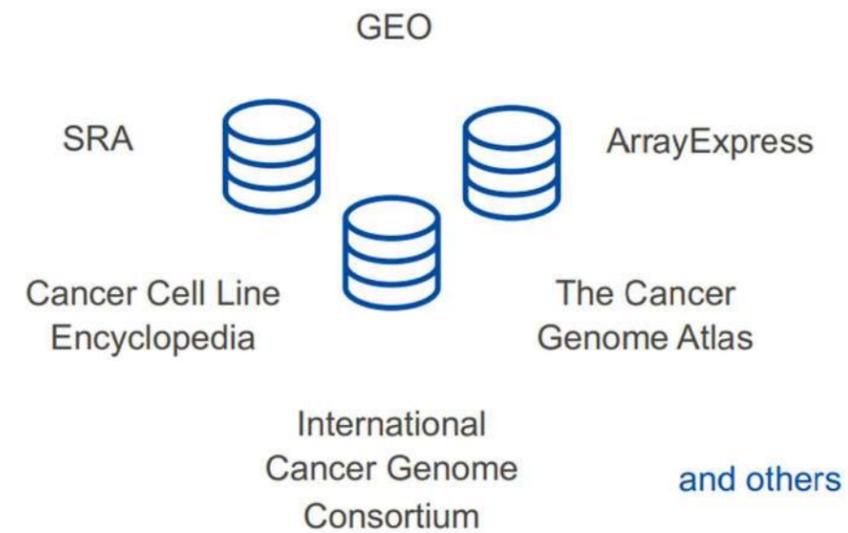
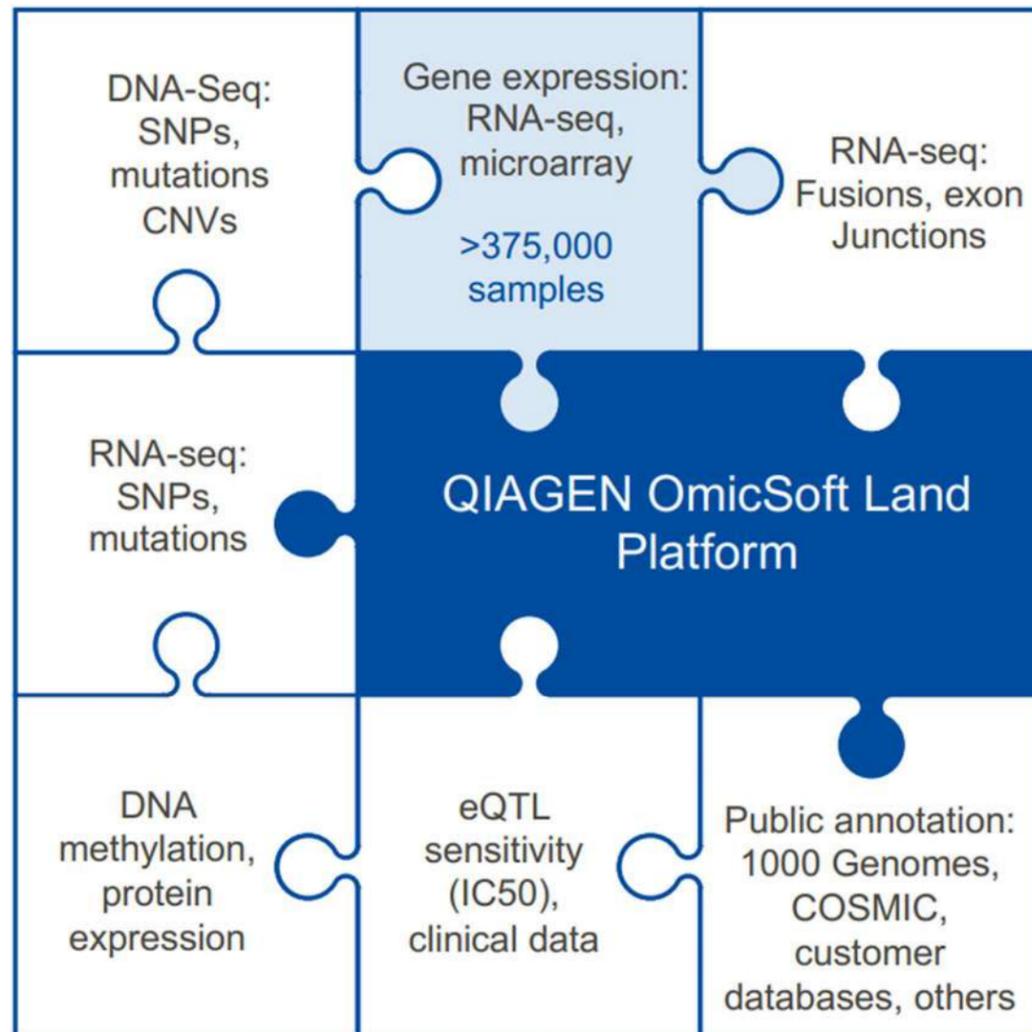






- More than **12 million** findings from literature source or database.
- More than **124,927** comparison datasets from QIAGEN OmicSoft Lands.

Lands provide expression data to QIAGEN IPA



Land	Repository	Datasets Q3 2022	Datasets Q4 2022	Increase
DiseaseLand	HumanDisease	31,041	32,092	1,051
	MouseDisease	24,411	24,506	95
	RatDisease	7,900	7,900	
	LINCS	28,234	28,234	
OncoLand	OncoHuman (Formerly OncoGEO)	13,360	14,131	771
	OncoMouse	1,054	1,054	
	TCGA	4,438	4,438	
	MetastaticCancer	81	81	
	Hematology	4,267	4,267	
	Pediatrics	444	444	
	ENCODE RNA Binding	486	486	
Single Cell Land	SingleCellHuman	194	194	
	SingleCellHumanUmi	2056	2056	
	SingleCellHumanUmiLite	603	603	
	SingleCellHumanHCL	1,476	1,476	
	SingleCellMouse	81	81	
	SingleCellMouseUmi	1,457	1,457	
	SingleCellMouseUmiLite	115	115	
Normal Cells and Tissues	Human Tissues (GTEx)	52	1,312	1,260

## Fully supported:



Human



Mouse



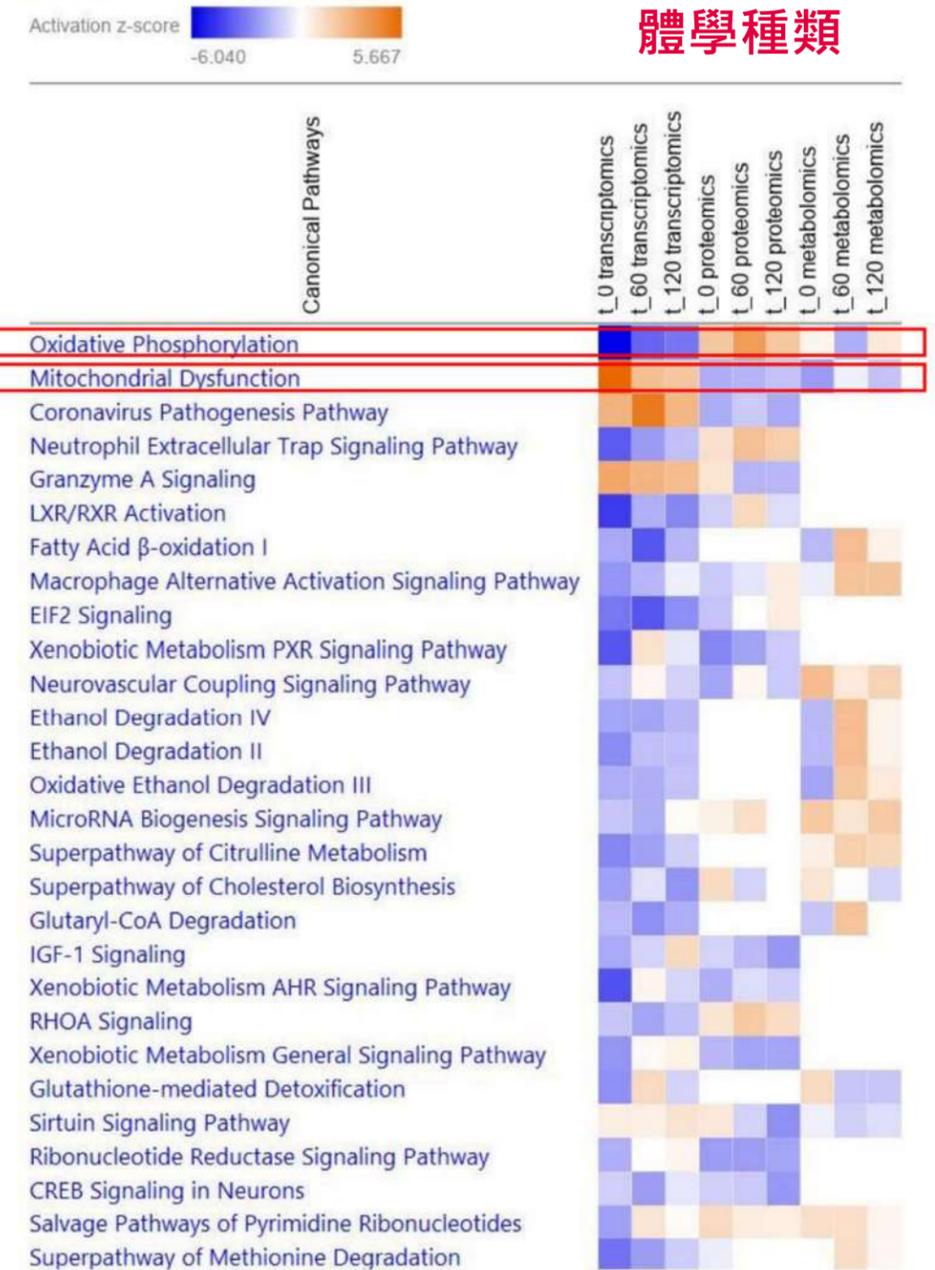
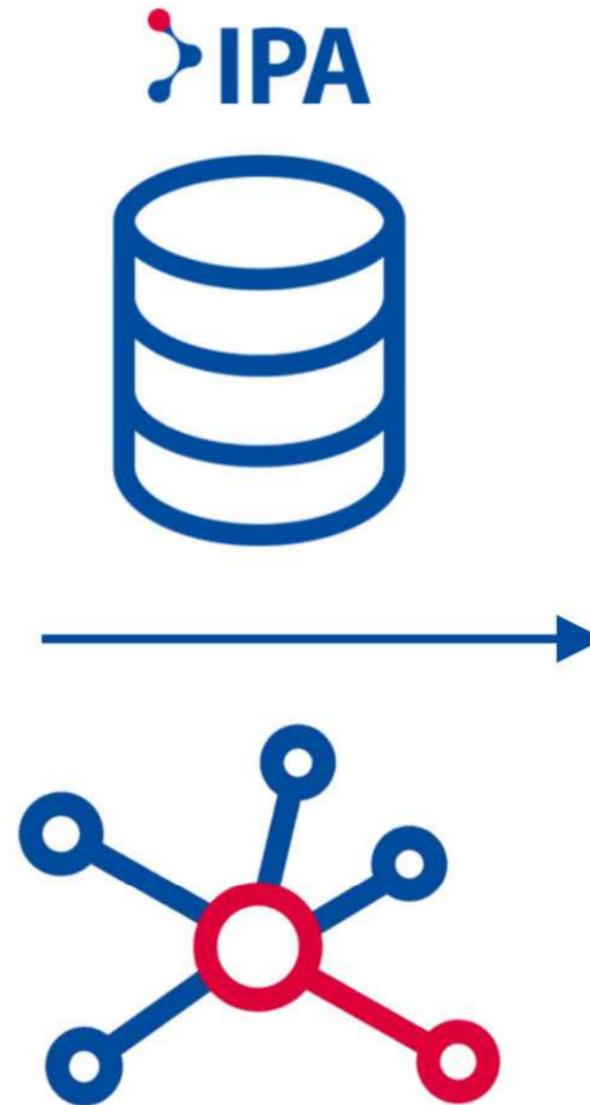
Rat

## IPA can remap orthologous identifiers from the following species:

- *Arabidopsis thaliana* (*Thale cress*)
- Bat (*Rhinolophus ferrumequinum*)
- *Caenorhabditis elegans*
- Cat (domestic, *Felis catus*)
- Chicken (*Gallus gallus*)
- Chimpanzee (*Pan troglodytes*)
- Chinese hamster (*Cricetulus griseus*, CHO-K1)
- Cow (*Bos taurus*)
- Crab-eating macaque (*Macaca fascicularis*)
- Dog (*Canis lupus familiaris*)
- Fruit fly (*Drosophila melanogaster*)
- Golden hamster (*Mesocricetus auratus*)
- Guinea pig, domestic (*Cavia porcellus*)
- Horse (*Equus caballus*)
- Human (*Homo sapiens*)
- Mouse (*Mus musculus*)
- Pig (*Sus scrofa*)
- Rabbit (*Oryctolagus cuniculus*)
- Rainbow trout (*Oncorhynchus mykiss*)
- Rat (*Rattus norvegicus*)
- Rhesus Monkey (*Macaca mulatta*)
- *Saccharomyces cerevisiae*
- *Schizosaccharomyces pombe*
- Western clawed frog (*Xenopus tropicalis*)
- Zebrafish (*Danio rerio*)

## Omics data type

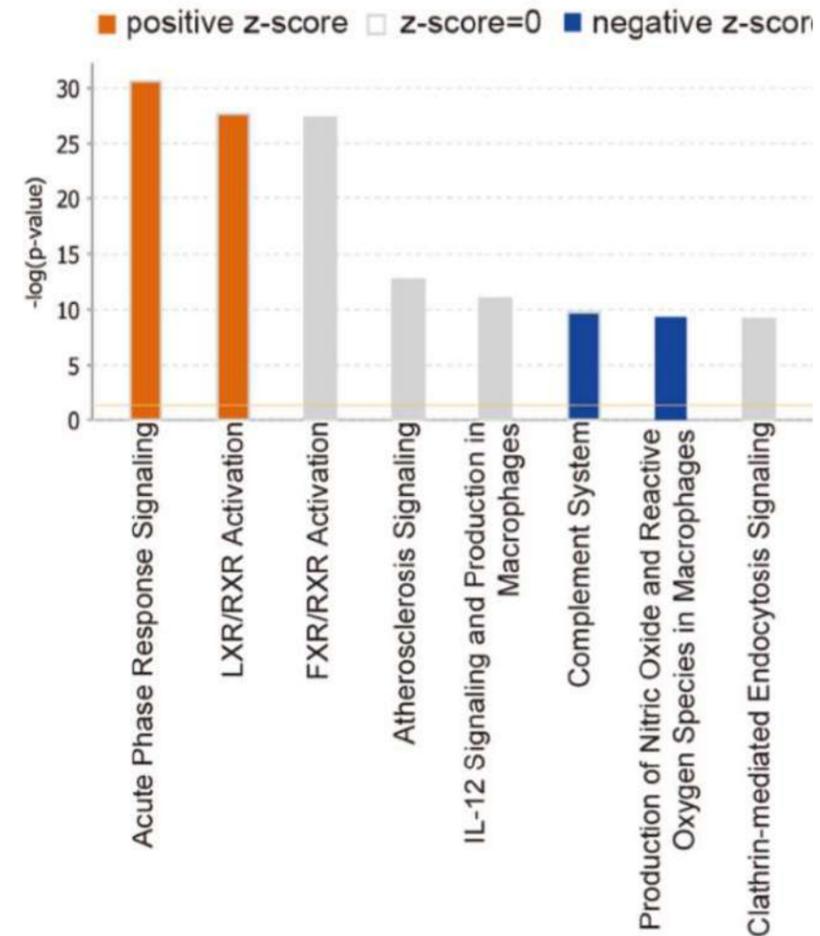
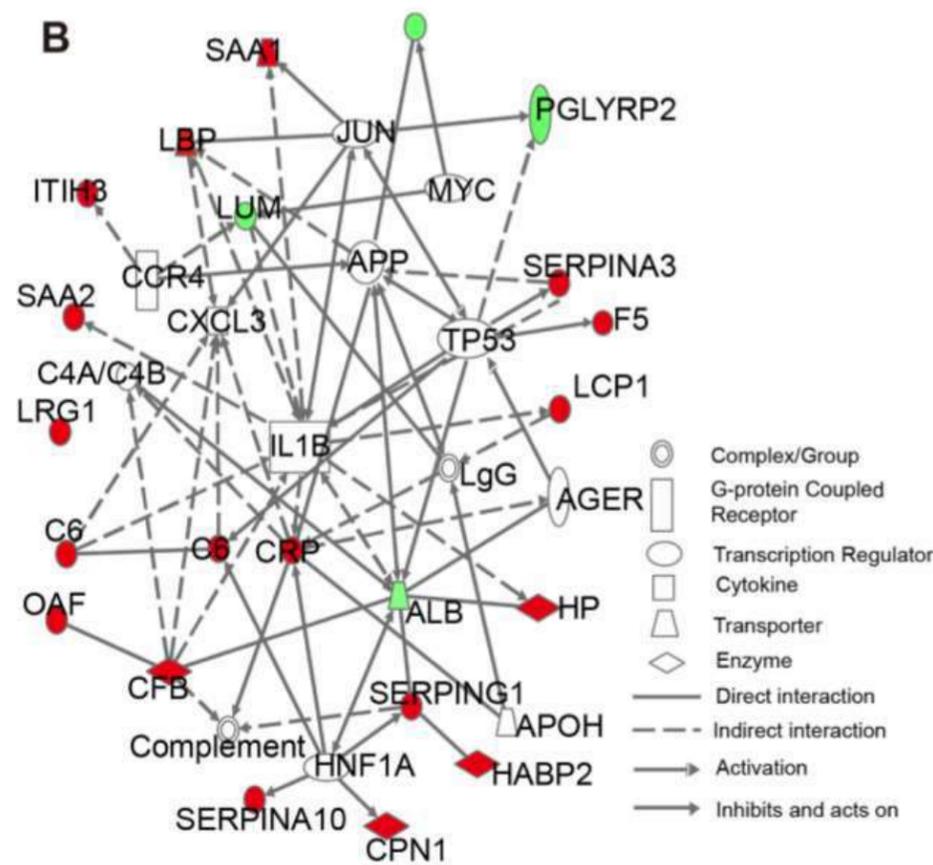
- RNA-seq
- scRNA-seq
- Microarray
- Nanostring
- qPCR
- ChIP-seq
- Proteomics
- Metabolomics
- RNAi
- CRISPR
- WGS/WES etc.



Get more complete mapping during dataset upload!

Vendor IDs	Gene	Protein	Transcript	microRNA	SNP	Chemical
Affymetrix (na36)	Entrez Gene (2020/10)	GenPept	Ensembl (101)	miRBase (mature)	Affy SNP IDs	CAS Registry Number
Agilent	GenBank (239)	International Protein Index (IPI)	RefSeq (human, mouse)	miRBase (stemloop)	dbSNP	HMDB
Life Tech (ABI)	Symbol-human (HUGO/ HGNC, EG)	UniProt/ Swiss-Prot Accession (2020_03)	UCSC (hg18)			KEGG
Codelink	Symbol- mouse (EG)		UCSC (hg19)			PubChem CID
Illumina	Symbol- rat (EG)		UCSC (hg38)			
Ingenuity	GI Number					
	UniGene					

Here is one latest paper submitted to medRxiv using IPA to identify potential protein and metabolite biomarkers in severe COVID-19 disease. In this study, IPA was used to identify most significantly relevant pathways and build regulation networks. It is welcome to share with your customers or distributor, and please let me know if you need further help.



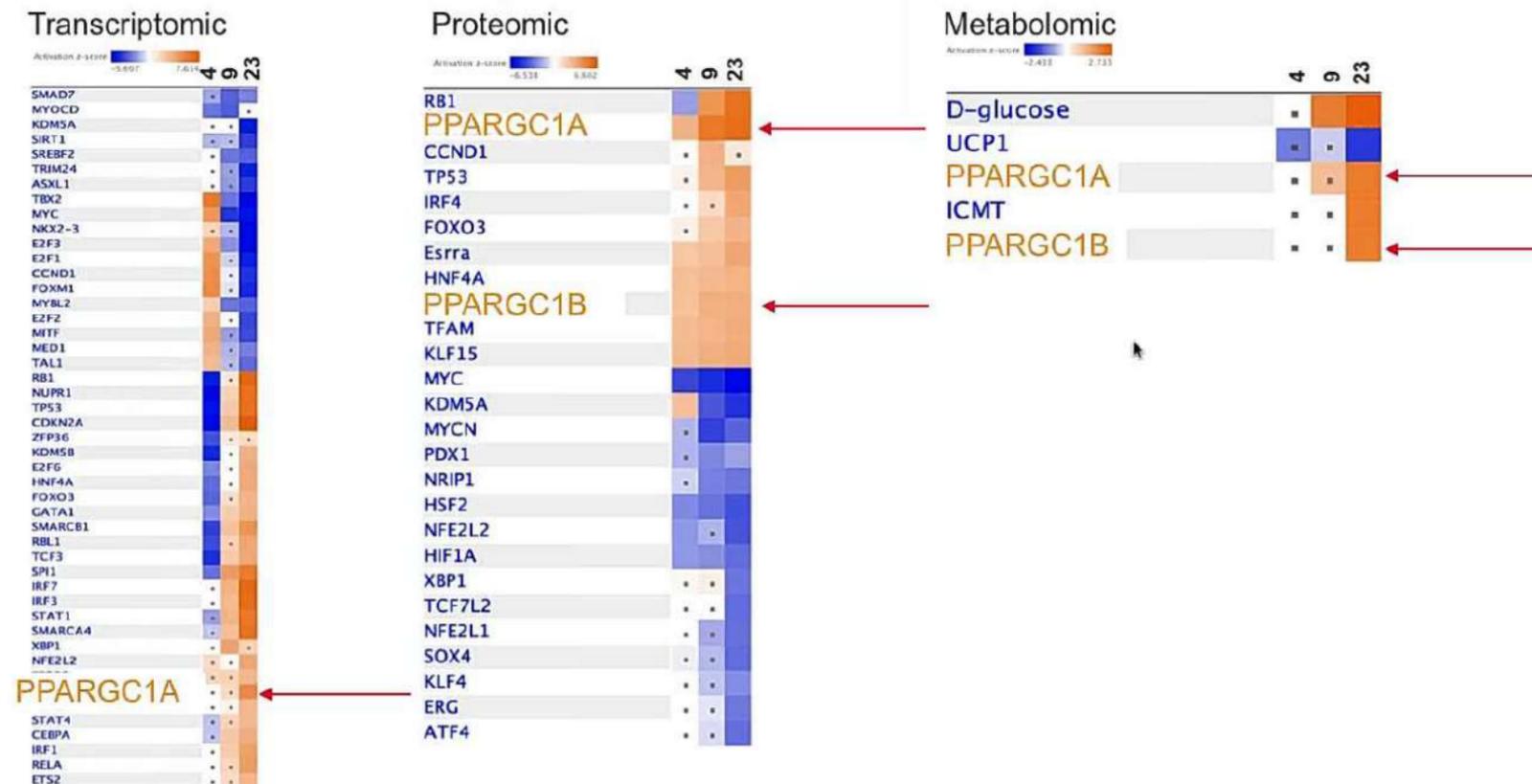
<https://www.medrxiv.org/content/10.1101/2020.04.07.20054585v1>

Integrate and compare genomics, transcriptomics, proteomics and metabolomics data to see the big picture on your focus research

Transcriptomics, proteomics and metabolic changes in the postnatal mouse heart identified by QIAGEN IPA and OmicSoft

Multi-omics analysis indicate similar transcriptional drivers

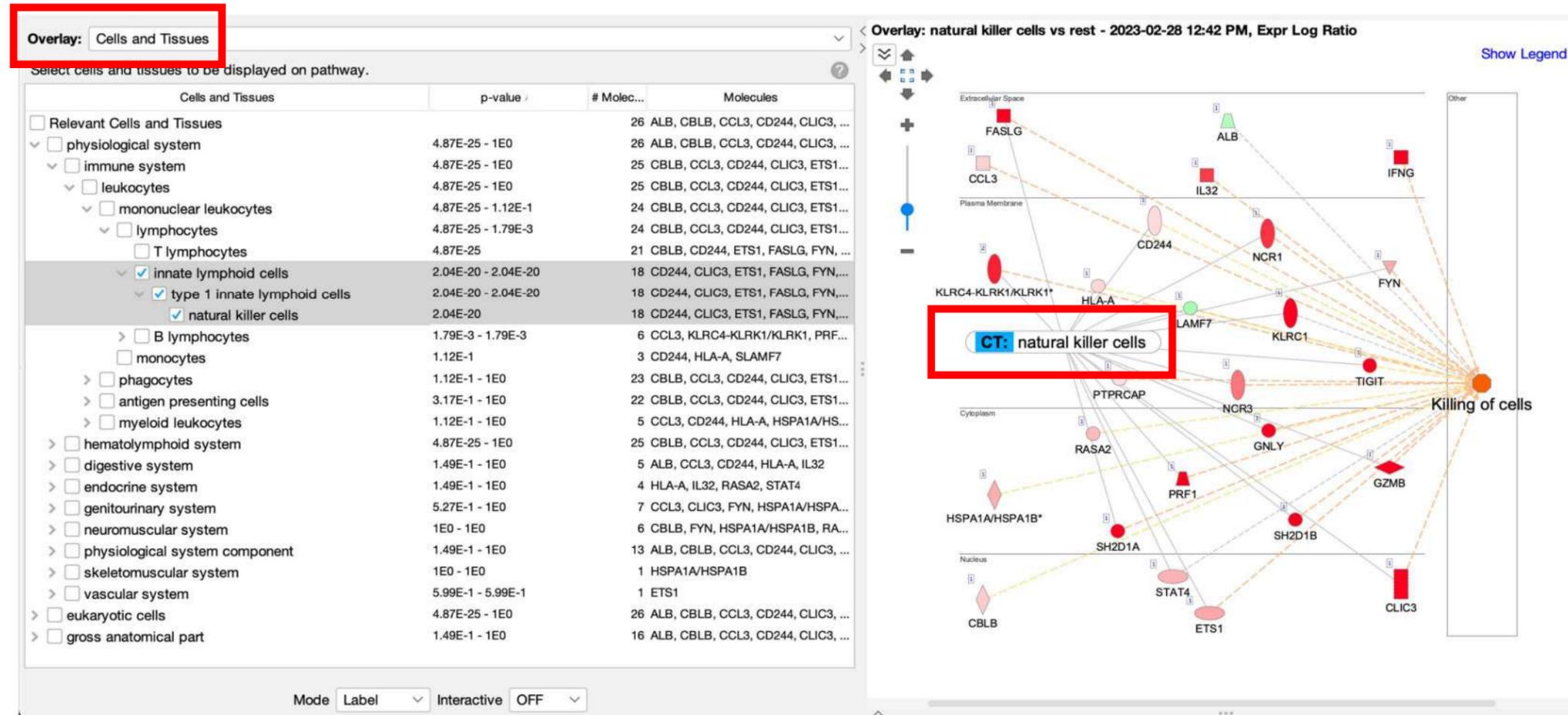
Upstream Analysis of transcriptomic, proteomic, and metabolomic data show induction of fatty oxidation regulation by PPARG coactivators.



<https://go.qiagen.com/IPA-multi-omics-analysis-webinar>

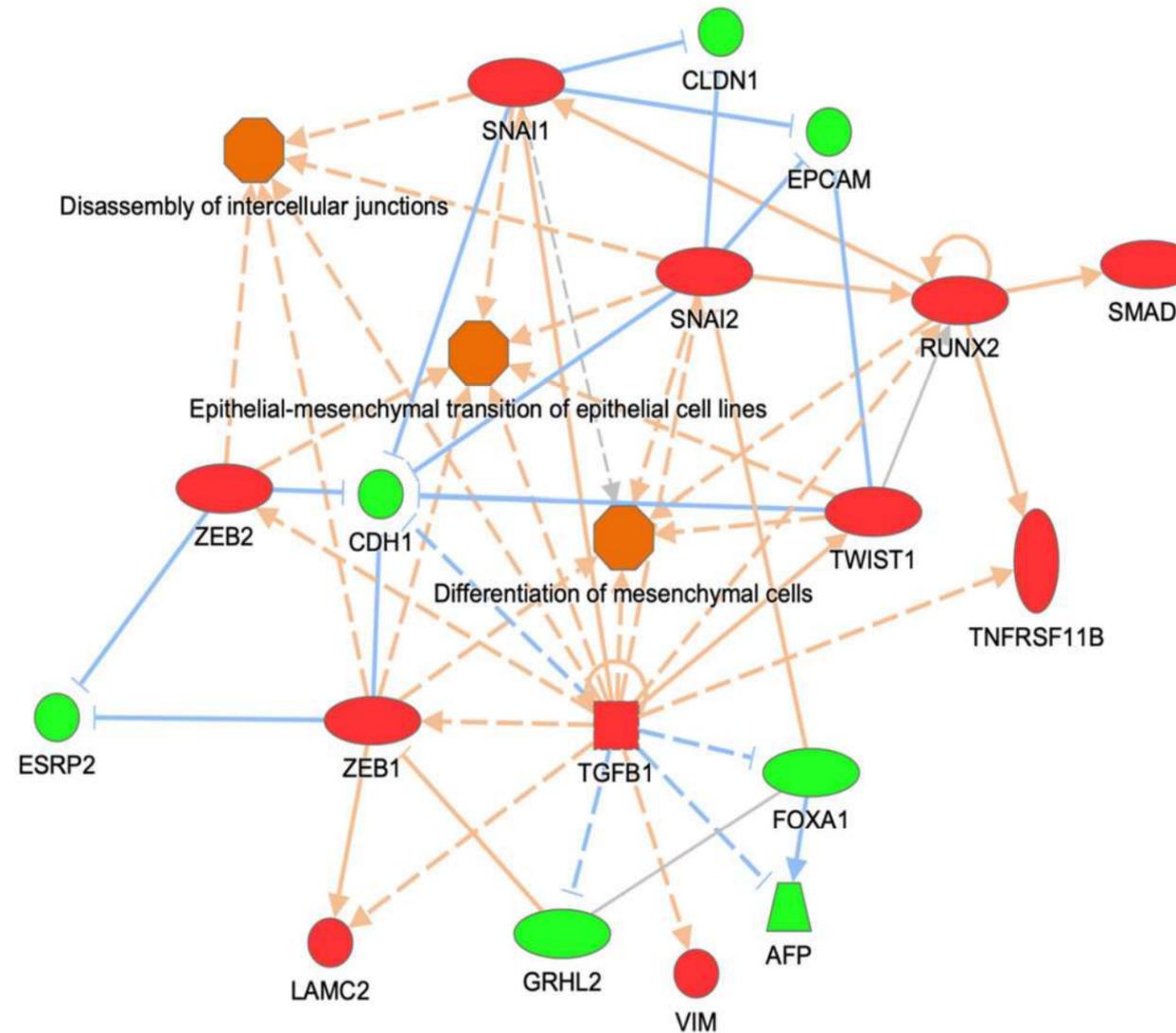
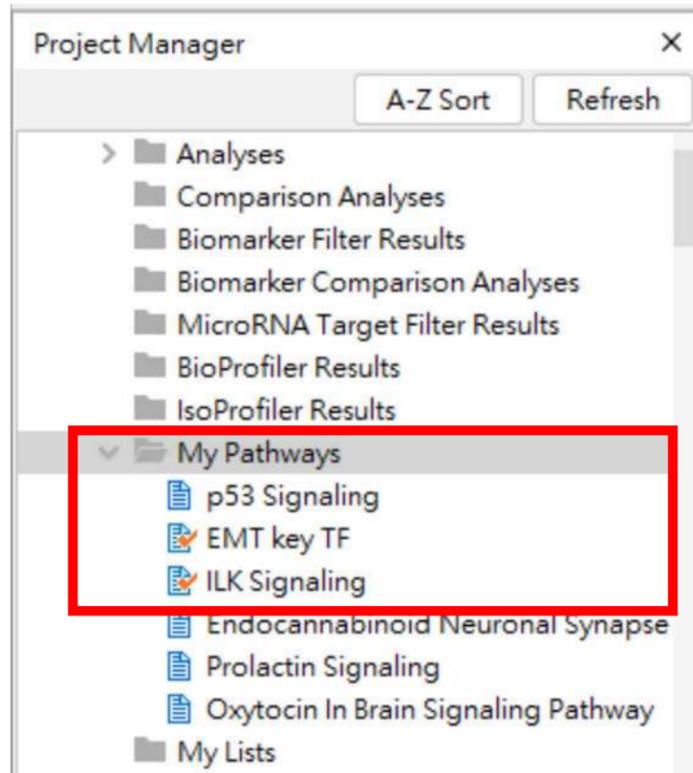
## Identify potential cell types based on the set of genes on networks and pathways

# Overlay :Cell and Tissues **THE HUMAN PROTEIN ATLAS**



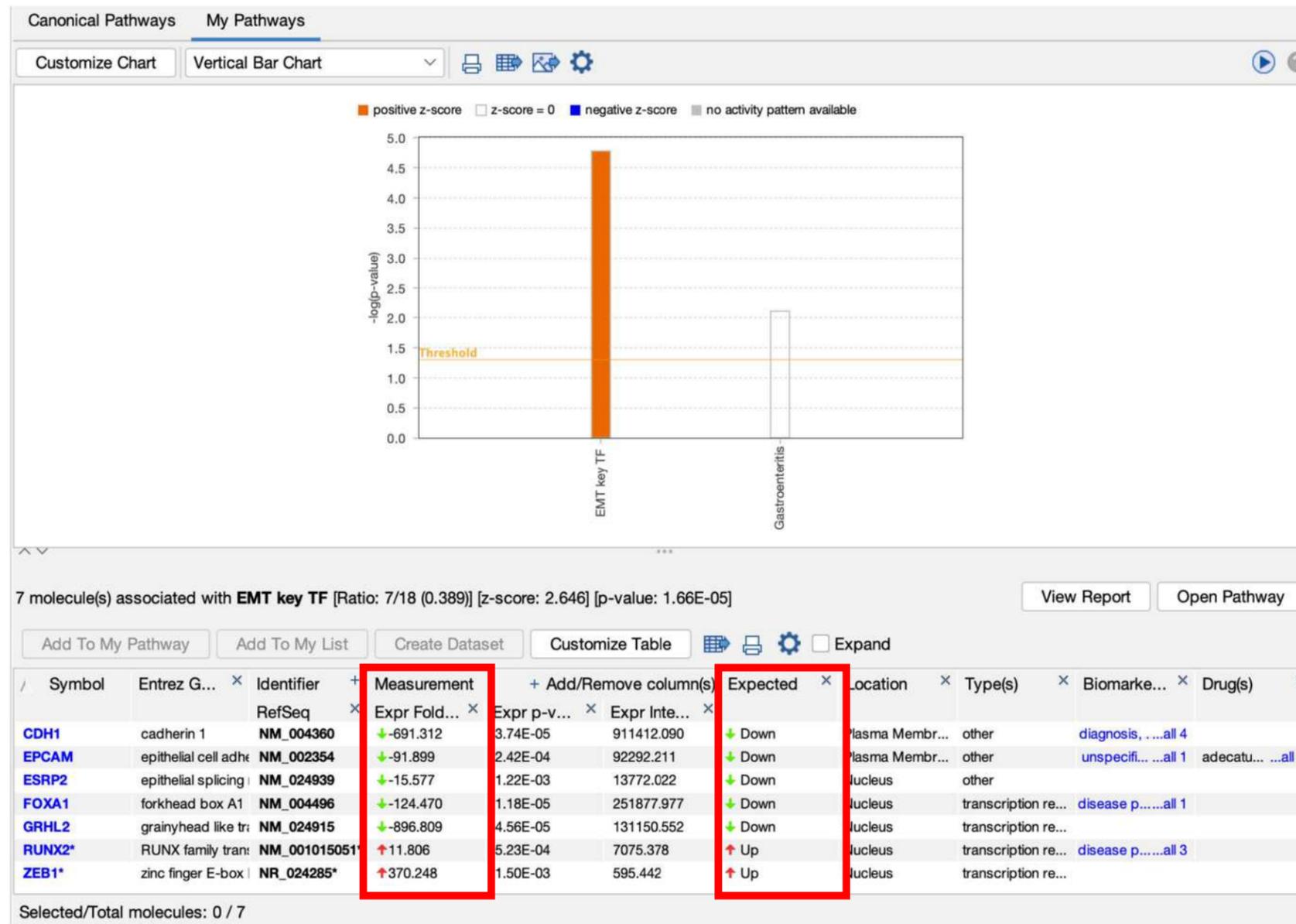
Enrichment of natural killer enriched genes on a network.

### Causally score My Pathways in Core Analysis



A custom My Pathway with nodes assigned by the user as activated (red) or green (inhibited).

## Causally score My Pathways in Core Analysis



Causally scoring a My Pathway.

## Set the “User Dataset” as the reference set when uploading a dataset

1. Select File Format: Flexible Format

2. Contains Column Header:  Yes  No

3. Select Identifier Type: Please assign at least one column below as “ID”, and assign additional columns as ID to improve mapping coverage.

4. Array platform used for experiments: Not specified/applicable

5. Use the dropdown menus to specify the reference set for the observations.

Raw Data (18589) Dataset Summary

Edit Observation Names Infer C

ID/Observation Name	ID	Measurement/Annotation	P-value	Bonferroni	FDR p-value
1	ID				
2	ENSMUSG000000051...		0.8708015979022331	1.0	0.9077575903
3	ENSMUSG000000025...		4.504633100231221...	8.57456910629013E...	4.4210170566
4	ENSMUSG000000033...		6.221800613621819...	1.184319746802913...	7.3009378079
5	ENSMUSG000000025...		1.682341354813838...	3.202336768888141...	1.4496821829

Setting the reference set to User Dataset reference during dataset upload.

## Discover more precise matches in Analysis Match

Summary Graphical Summary Pathways Upstream Analysis Diseases & Functions Regulator Effects Networks Lists Analysis Match Molecules

Evaluate Metadata View As Heatmap View Comparison Customize Table z-sc... 54.42 - 33.52 (1/539)

Analysis ...	Pro...	case....	case.t...	compar...	comparisoncontrast	T x	T x	T x	T x	T x	T x	T x
86- NA [heart] NA	MouseDise...	NA	heart	CellType1 vs. Ce...	CellDescription => heart epithelial cell vs cecum epithelial cell	67.94	50.99	45.83	52.92	54.42	35.33	
828- normal contr	MouseDise...	normal control	fetal heart	Other Compariso...	ExperimentGroup => fetal heart E11.5 vs fetal brain E11.5	62.02	56.57	46.90	50.99	54.12	49.48	
11- normal contro	HumanDise...	normal control	skin	CellType1 vs. Ce...	CellType => iPSC-derived cardiomyocyte vs induced pluripotent stem cell (iPSC)	55.47	61.64	44.72	44.72	51.64	30.88	
4- psoriasis [skin]	HumanDise...	psoriasis	skin	Tissue1 vs. Tissu...	TissueRegion => back vs arm	55.47	52.92	33.17	56.57	49.53	26.34	
150- normal contr	RatDisease...	normal control	heart	Tissue1 vs. Tissu...	Tissue:Age[weeks] => 21 -> heart vs testis	55.47	45.83	50.00	45.83	49.28	34.66	
30- normal contro	Human Tiss...	normal control	heart atria	Other Compariso...	TissueDetail_GTEx => Heart - Atrial Appendage vs Others	62.02	46.90	37.42	50.00	49.08	30.12	
1856- normal con	MouseDise...	normal control	pancreas	Other Compariso...	Tissue:Age[months] => pancreas -> 27 vs 24	62.02	44.72	42.43	46.90	49.02	33.09	
132- normal contr	RatDisease...	normal control	heart	Tissue1 vs. Tissu...	Tissue:Age[weeks] => 104 -> heart vs testis	55.47	41.23	50.00	47.96	48.66	35.50	
13- fibromyxoid s	OncoHuman	fibromyxoid sa...	soft tissue	Disease1 vs. Dis...	ExperimentGroup => fibromyxoid sarcoma vs Castleman disease follicular dend...	73.38	50.99	25.30	40.00	47.42	17.08	
31- normal contro	Human Tiss...	normal control	heart left ventricle	Other Compariso...	TissueDetail_GTEx => Heart - Left Ventricle vs Others	55.47	45.83	37.42	48.99	46.93	29.94	

Sorted by original overall z-score match

Matches to analyses from many tissue types, such as skin and pancreas

Summary Graphical Summary Pathways Upstream Analysis Diseases & Functions Regulator Effects Networks Lists Analysis Match Molecules

Evaluate Metadata View As Heatmap View Comparison Customize Table DM (... 52.1226215454257... (1/539)

Analysis ...	Pro...	case....	case.t...	compar...	comparisoncontrast	T x	T x	T x	T x	T x	T x	T x
3- normal control	MouseDise...	normal control	hindlimb muscle	CellType1 vs. Ce...	SampleDescription => primary myotubes differentiated for 1 day vs embryonic ...	46.90	33.17	30.00	27.52	52.12		
2- normal control	MouseDise...	normal control	hindlimb muscle	CellType1 vs. Ce...	SampleDescription => primary myoblasts passage zero vs embryonic stem cell	43.59	41.23	40.00	31.21	51.80		
5- normal control	MouseDise...	normal control	hindlimb muscle	CellType1 vs. Ce...	SampleDescription => primary myotubes differentiated for 4 days vs embryonic...	44.26		30.51	18.69	50.70		
1- normal control	MouseDise...	normal control	hindlimb muscle	CellType1 vs. Ce...	SampleDescription => primary myoblasts in proliferation passage 8-10 vs embr...	43.59	31.62	38.73	28.49	49.71		
828- normal contr	MouseDise...	normal control	fetal heart	Other Compariso...	ExperimentGroup => fetal heart E11.5 vs fetal brain E11.5	62.02	56.57	46.90	50.99	54.12	49.48	
6- normal control	MouseDise...	normal control	quadriceps fem.	Other Compariso...	SampleDescription => quadriceps muscle vs embryonic stem cell	28.40			32.07	15.12	49.35	
4- normal control	MouseDise...	normal control	hindlimb muscle	CellType1 vs. Ce...	SampleDescription => primary myotubes differentiated for 2 days vs embryonic...	55.47	39.62		28.87	30.99	49.35	
11- normal contro	MouseDise...	normal control	embryo	CellType1 vs. Ce...	CellType => ESC-derived NPC vs embryonic stem cell (ESC)	37.42	31.62	31.62	25.17	49.16		
843- normal contr	MouseDise...	normal control	fetal heart	Other Compariso...	ExperimentGroup => fetal heart E12.5 vs fetal brain E12.5	55.68	45.83	23.24	31.19	45.43		
988- normal contr	RatDisease...	normal control	fetal heart	Other Compariso...	ExperimentGroup => fetal heart E11 vs fetal brain E11	55.68	38.73	33.00	31.85	45.34		

Sorted by new DM "dataset match" column

Results are more strictly heart or muscle cells vs embryonic cells

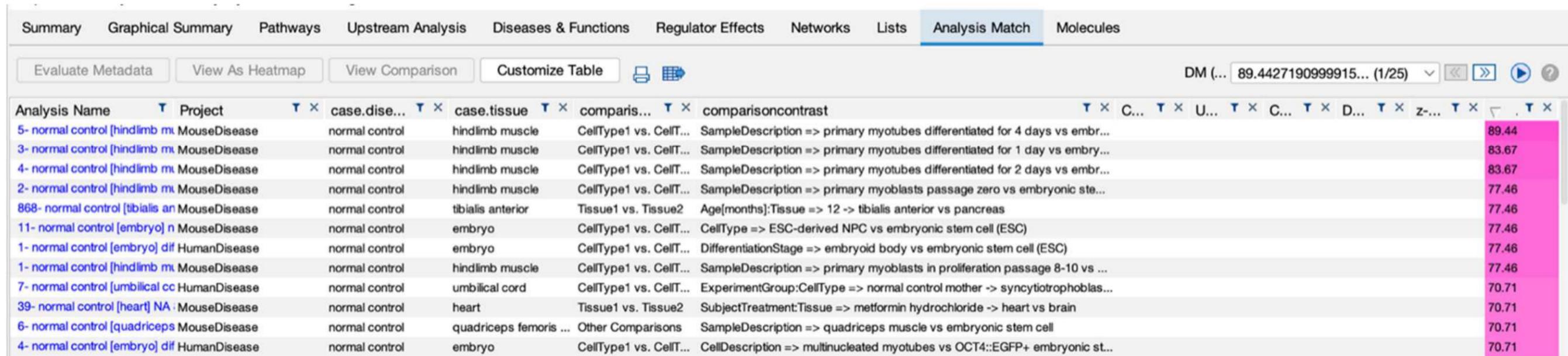
Analysis Match results sorted by the original score (top) and new score (bottom).





## The new scoring method using a small dataset.

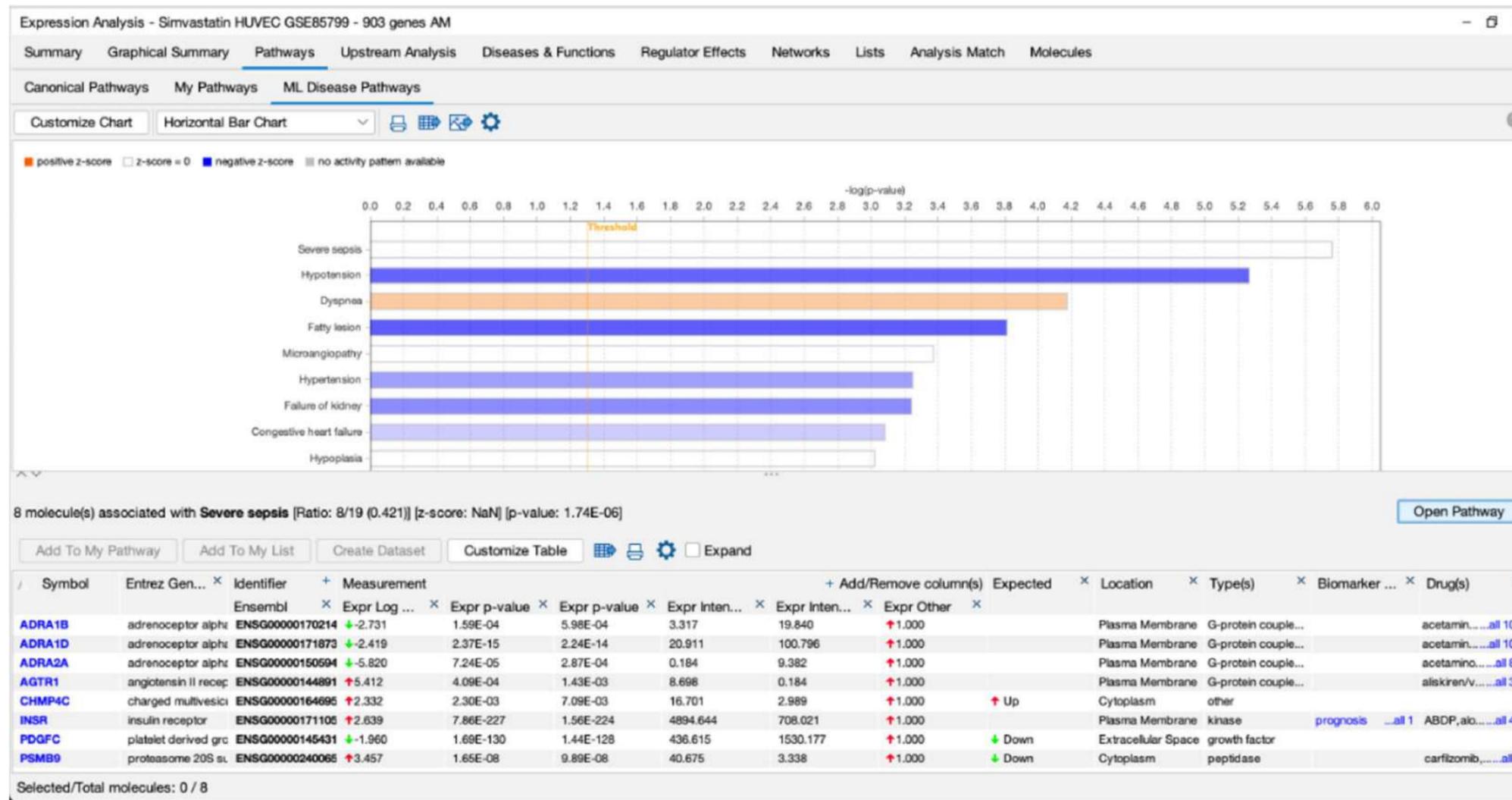
The new scoring method often works on small datasets, where there are typically too few genes to generate robust Upstream Regulator, Causal Network, Canonical Pathway, and Disease and Function signatures to match to other analyses.



Analysis Name	Project	case.dise...	case.tissue	comparis...	comparisoncontrast	Score
5- normal control [hindlimb m]	MouseDisease	normal control	hindlimb muscle	CellType1 vs. CellT...	SampleDescription => primary myotubes differentiated for 4 days vs embr...	89.44
3- normal control [hindlimb m]	MouseDisease	normal control	hindlimb muscle	CellType1 vs. CellT...	SampleDescription => primary myotubes differentiated for 1 day vs embry...	83.67
4- normal control [hindlimb m]	MouseDisease	normal control	hindlimb muscle	CellType1 vs. CellT...	SampleDescription => primary myotubes differentiated for 2 days vs embr...	83.67
2- normal control [hindlimb m]	MouseDisease	normal control	hindlimb muscle	CellType1 vs. CellT...	SampleDescription => primary myoblasts passage zero vs embryonic ste...	77.46
868- normal control [tibialis an]	MouseDisease	normal control	tibialis anterior	Tissue1 vs. Tissue2	Age[months]:Tissue => 12 -> tibialis anterior vs pancreas	77.46
11- normal control [embryo] n	MouseDisease	normal control	embryo	CellType1 vs. CellT...	CellType => ESC-derived NPC vs embryonic stem cell (ESC)	77.46
1- normal control [embryo] dif	HumanDisease	normal control	embryo	CellType1 vs. CellT...	DifferentiationStage => embryoid body vs embryonic stem cell (ESC)	77.46
1- normal control [hindlimb m]	MouseDisease	normal control	hindlimb muscle	CellType1 vs. CellT...	SampleDescription => primary myoblasts in proliferation passage 8-10 vs ...	77.46
7- normal control [umbilical cc]	HumanDisease	normal control	umbilical cord	CellType1 vs. CellT...	ExperimentGroup:CellType => normal control mother -> syncytiotrophoblas...	70.71
39- normal control [heart] NA	MouseDisease	normal control	heart	Tissue1 vs. Tissue2	SubjectTreatment:Tissue => metformin hydrochloride -> heart vs brain	70.71
6- normal control [quadriceps]	MouseDisease	normal control	quadriceps femoris ...	Other Comparisons	SampleDescription => quadriceps muscle vs embryonic stem cell	70.71
4- normal control [embryo] dif	HumanDisease	normal control	embryo	CellType1 vs. CellT...	CellDescription => multinucleated myotubes vs OCT4::EGFP+ embryonic st...	70.71

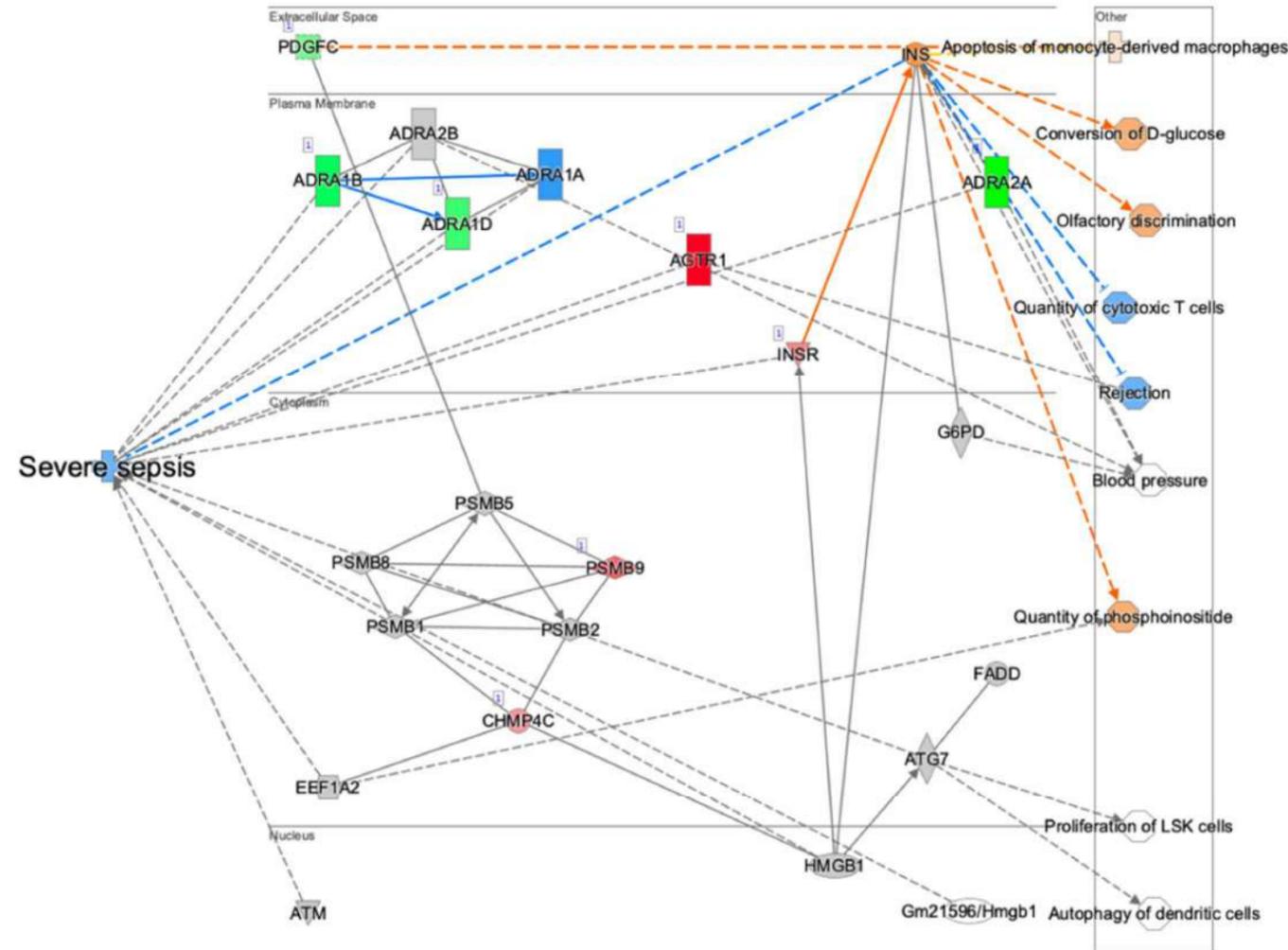
The analysis of a 10-gene dataset from the cardiomyocytes versus embryonic stem cells matches the expected types of analyses.

## Find the unexpected with ML Disease Pathways in Core Analyses



ML Disease Pathways scored against simvastatin-treated rats (liver). The most significant result by Fisher's Exact Test (right-tailed) is "Severe sepsis".

## Find the unexpected with ML Disease Pathways in Core Analyses

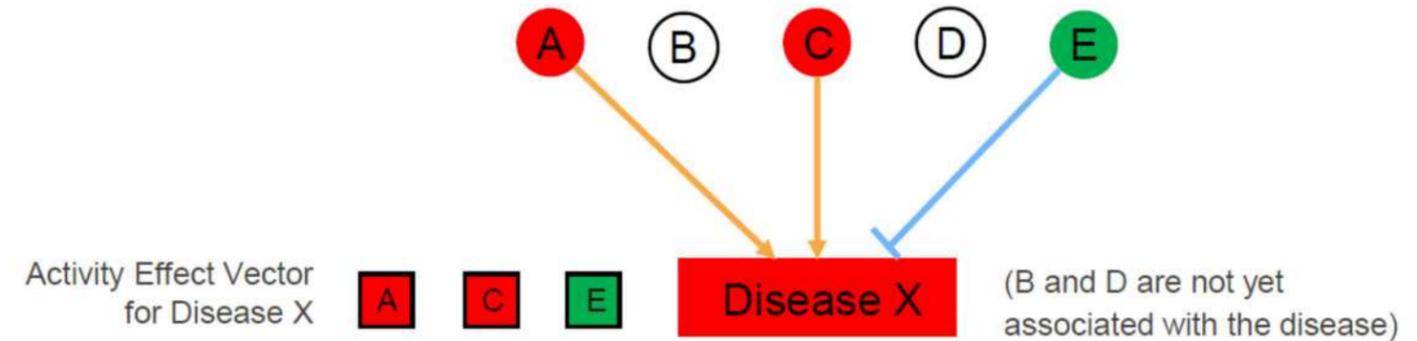


The severe sepsis ML pathway overlaid with simvastatin differential expression data. IPA predicts that simvastatin may decrease severe sepsis.

## Enhancing the KB through Machine Learning

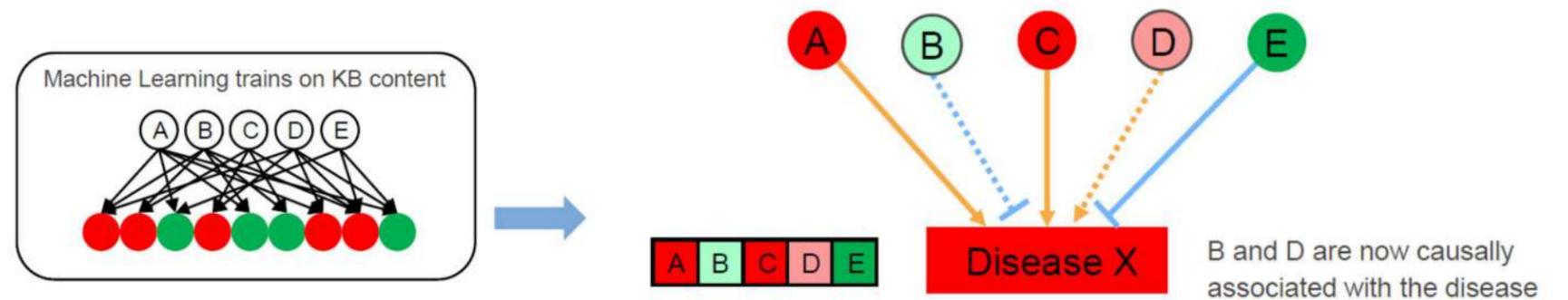
### Standard KB

- Relationships between Disease X and neighboring genes can be described as an Activity Effect Vector built only from KB findings.



### Enhanced KB

- Machine Learning builds richer Activity Effect Vectors by identifying molecules that have similar activity effects. Such vectors enable the inference of new associations and prioritization of relationships.
- In short, if two genes regulate the expression of a similar set of genes, but only one is associated with the disease, then the other likely has a similar biological impact as well.



Activity Effect Vectors consist of thousands of molecules in the KB. The approach has been validated by “rediscovering” known, curated findings in the KB.

## Other software changes

- The speed to upload and save a dataset has been improved, most noticeably for datasets with many columns.
- “Reactome” appears as a content source in certain filters in the UI in this release, however, there is currently only a minor addition of Reactome content in IPA. As we prepare to add Reactome pathways in a future release, that source currently only refers to new groups and complexes that have added from Reactome.

## New pathways

- Acetylcholine Receptor Signaling Pathway
- Adrenergic Receptor Signaling Pathway (Enhanced)
- Cachexia Signaling Pathway
- GABAergic Receptor Signaling Pathway (Enhanced)
- Glutaminergic Receptor Signaling Pathway (Enhanced)
- ISGylation Signaling Pathway
- Microautophagy Signaling Pathway
- NFKBIE Signaling Pathway
- Orexin Signaling Pathway
- Sertoli Cell Germ Cell Junction Signaling Pathway (Enhanced)

## Existing pathways updated to include an activity pattern

- IL-17A Signaling in Fibroblasts
- Sertoli Cell-Sertoli Cell Junction Signaling
- TR/RXR Activation

## **Addition of >400,000 new findings (bringing the total in IPA to over 12.6 million)**

- >29,000 protein-protein interaction findings from BioGrid
- >407,000 cancer mutation findings from ClinVar
- >1,800 target-to-disease findings from ClinicalTrials.gov
- >1,700 drug-to-disease findings from ClinicalTrials.gov
- >800 Gene Ontology findings
- >220 mappable chemicals
- > 3,800 Lipid Maps IDs

## 141,323 expression datasets (5,689 added)

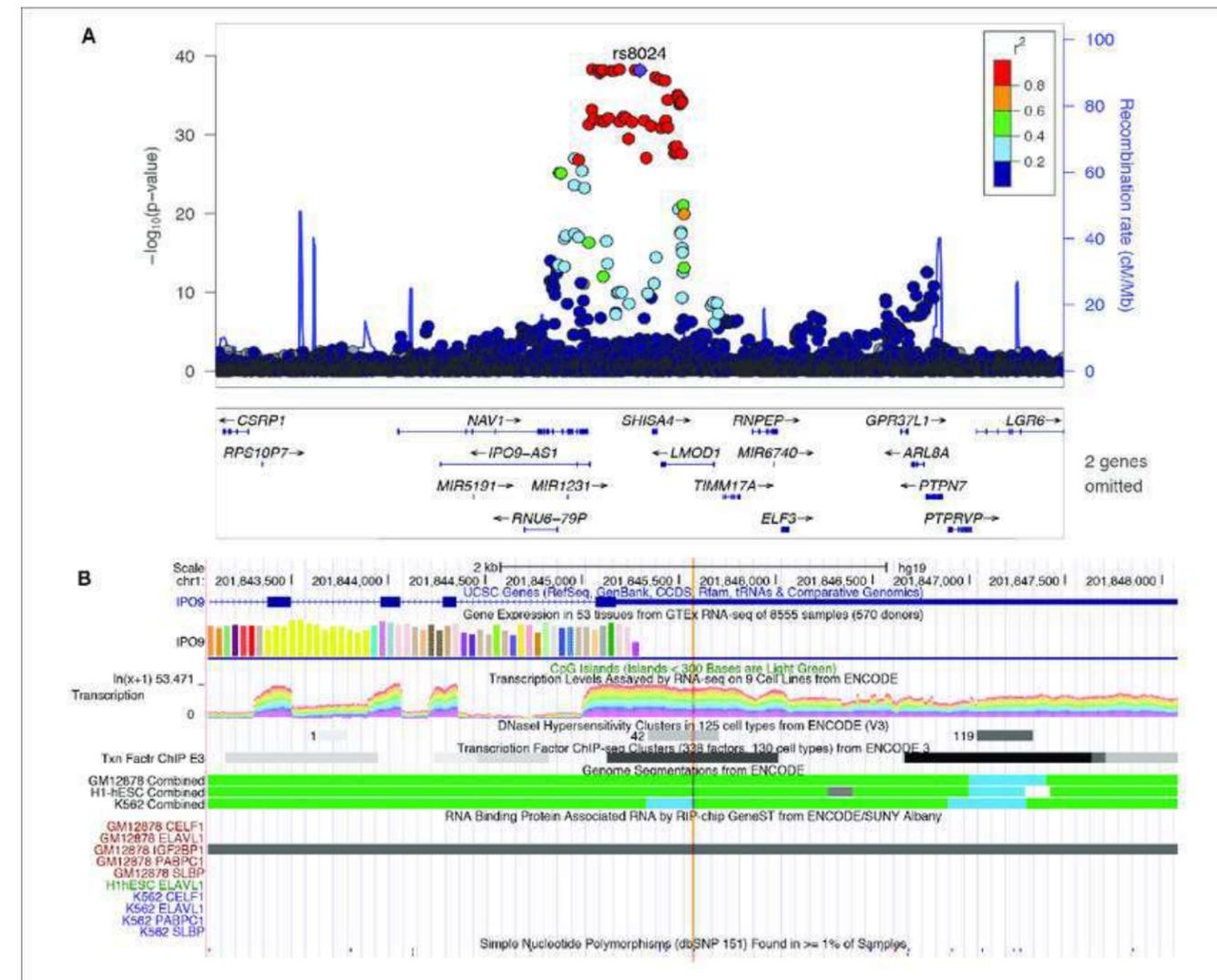
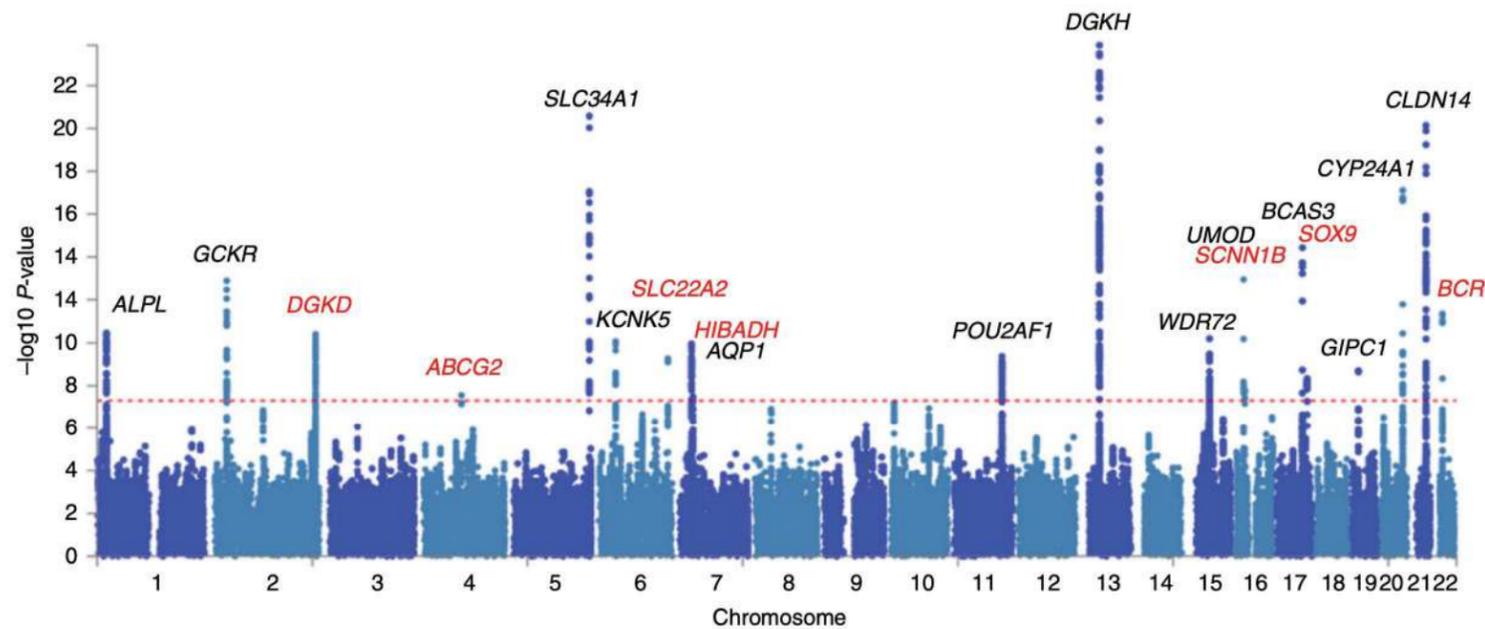
Land	Repository	Datasets Q1 2023	Datasets Q2 2023	Increase
DiseaseLand	HumanDisease	32,680	33,672	992
	MouseDisease	25,070	25,253	183
	RatDisease	8,226	8,226	
	LINCS	28,234	28,234	
OncoLand	OncoHuman (Formerly OncoGEO)	15,147	17,125	1,978
	OncoMouse	1,054	1,054	
	TCGA	4,438	4,438	
	MetastaticCancer	81	81	
	Hematology	4,267	4,267	
	Pediatrics	444	444	
	ENCODE RNA Binding*	486	486	
Single Cell Land	SingleCellHuman	194	194	
	SingleCellHumanUmi	8,636	11,049	2,413
	SingleCellHumanUmiLite	603	603	
	SingleCellHumanHCL	1,476	1,469	-7
	SingleCellMouse	81	81	
	SingleCellMouseUmi	3,097	3,220	123
	SingleCellMouseUmiLite	115	115	
Normal Cell and Tissues	Human Tissues (GTEx)	1,312	1,312	

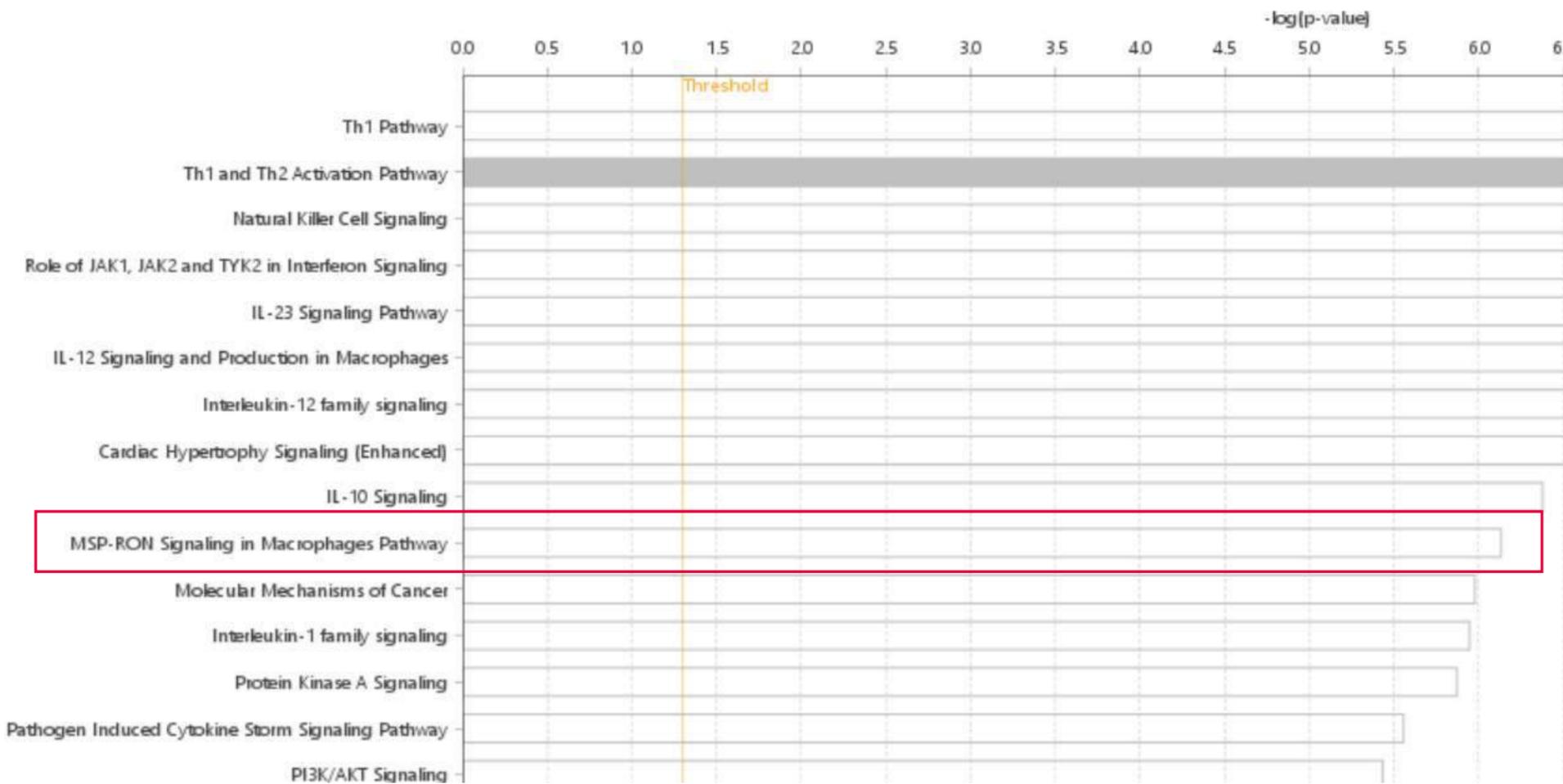
	IPA	Metacore	KEGG	Note
<b>資料檢索</b>				
基因、蛋白質	○	○	○	
藥物、毒物、代謝物	○	○	○	
疾病與生物功能	○	○	○	
Pathway	○	○	○	
miRNA	○	○	○	
調控作用關係	○	○	○	
Isoform	○	X	X	
Single Cell DB	○	X	X	
Patient Metadata	○	X	X	
Clinical data	○	X	X	
<b>繪製Pathway功能</b>				
互動式Pathway繪製工具	○	○	X(僅提供Mapping)	
Pattern Mapping	○	X	X	根據產生出來的Network以及實驗表現結果，搜尋資料庫疾病、Pathway是否有類似的表現趨勢以及路徑。
Path Tracer	○	X	X	在龐大的訊息路徑中，特別標示感興趣的訊息因子。
表現預測功能	○	○	X	根據實驗結果，預測上下游調控因子表現。

	IPA	Metacore	KEGG	Note
<b>Enrichment Analysis</b>				
基因體、蛋白質體表現分析	○	○	○	
代謝體	○	○	○	
毒物分析	○	○	○	
Enrichment Analysis結果視覺化表現	○	X	X	將各項Enrichment結果以不同的視覺化分析圖表呈現。
Enrichment Analysis Expression Pattern比較	○	X	X	將本次Enrichment結果與其他疾病、實驗的Dataset進行相似性比較。
因果網路分析	○	X	X	將Enrichment結果與感興趣的生物分子或者疾病、生物功能產生關連。
疾病熱圖	○	X	X	以Ontology分類，依據Enrichment分析p-value進行熱圖繪製，可找出較相關的生物功能以及疾病。
網絡關聯圖	○	X	X	將各項Enrichment結果依照重疊的訊息傳遞因子數量進行關聯統計。
泡泡圖分析	○	X	X	以泡泡圖進行三維分析數據結果解讀。可依照生物功能、疾病或者訊息路徑比對關聯性。
比較結果熱圖	○	X	X	以熱圖表示不同Enrichment結果差異，可用做Time or dosage – dependent的實驗比較。

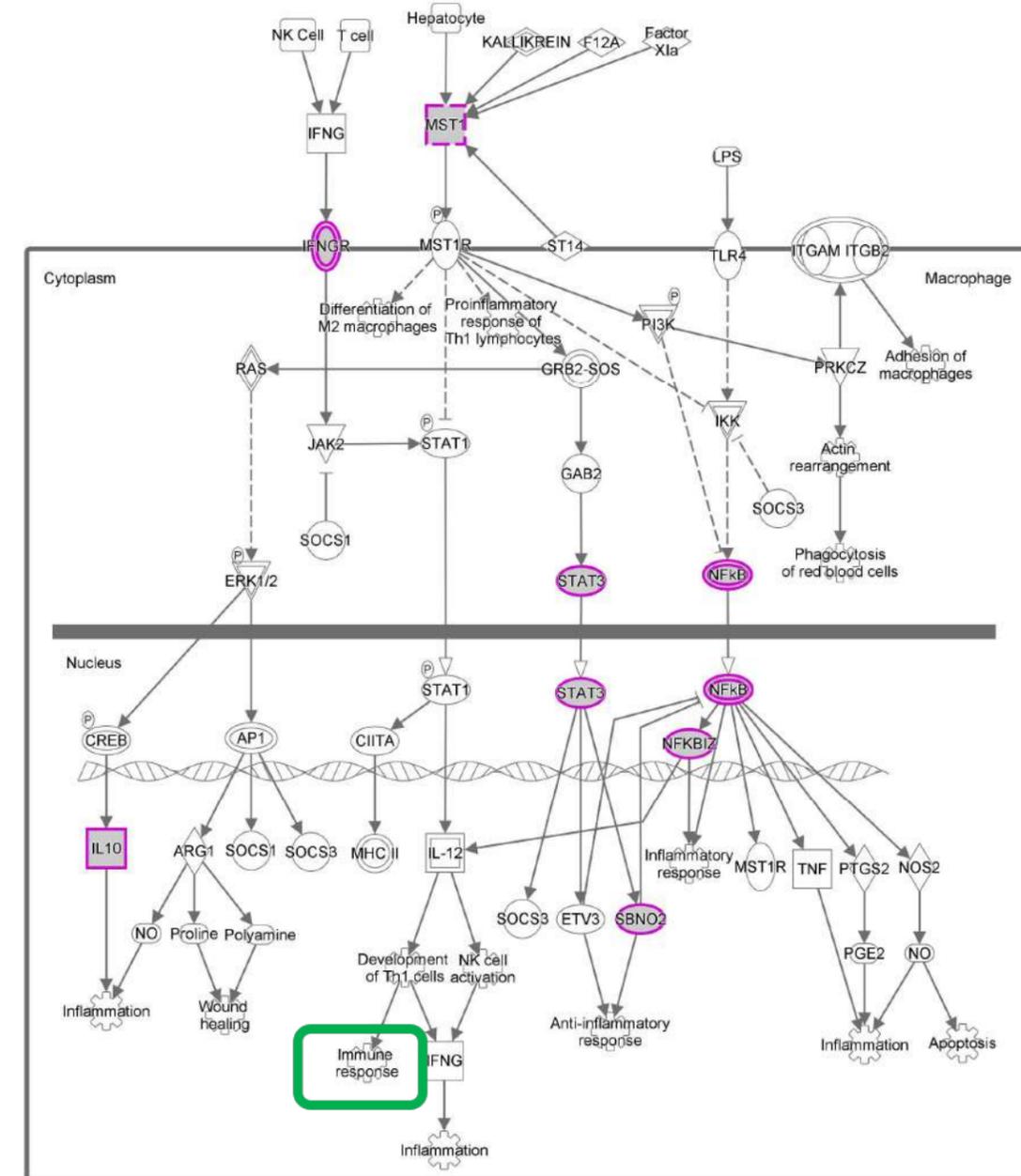
	IPA	Metacore	KEGG	Note
<b>其他功能</b>				
生物特性篩選	○	X	X	Bioprofiler，可篩選出具有相同生物特性的訊息傳遞因子。
Isoform篩選	○	X	X	Isoprofiler，可鑑別出同基因不同Isoform的生物表現結果。
基因於組織差異表現分析	○	X	X	GTEX Expression Pattern，用作於比較相同基因在不同組織的差異分析。
miRNA調控目標預測分析	○	X	X	miRNA Target filter，可進行miRNA與mRNA配對，進而比較出miRNA的作用方式。
磷酸化蛋白質體分析	○	X	X	用於確認不同位點的磷酸化蛋白質功能。
AI預測訊息路徑	○	○ (僅有Drug)	X	透過機器學習方式，搜尋相似表現路徑的訊息傳遞因子，連結其相關影響疾病和生物功能。

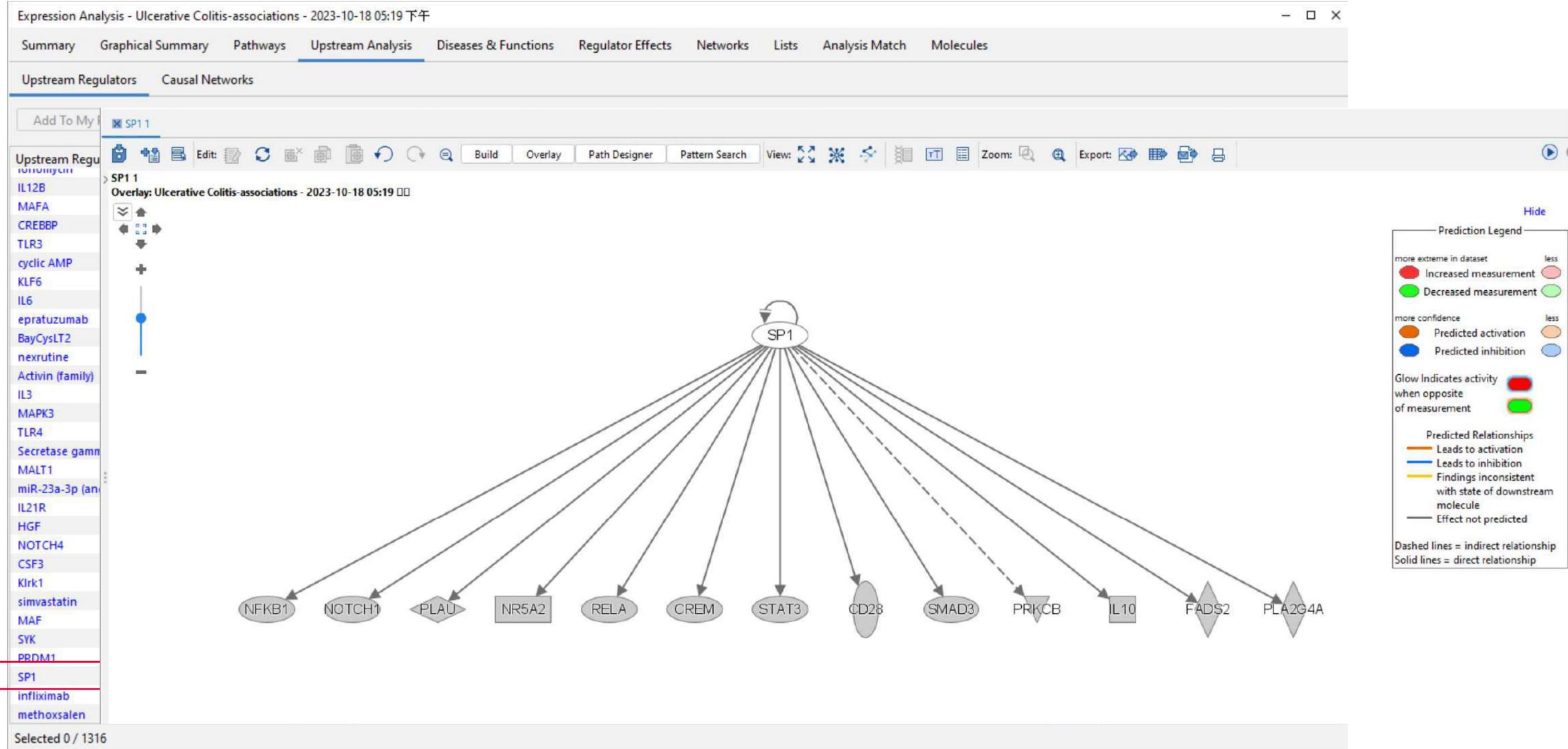
A genome-wide association study (GWA study, or GWAS), is an observational study of a genome-wide set of genetic variants in different individuals to see if any variant is associated with a trait. GWAS typically focus on associations between single-nucleotide polymorphisms (SNPs) and traits like major human diseases, but can equally be applied to any other genetic variants and any other organisms.



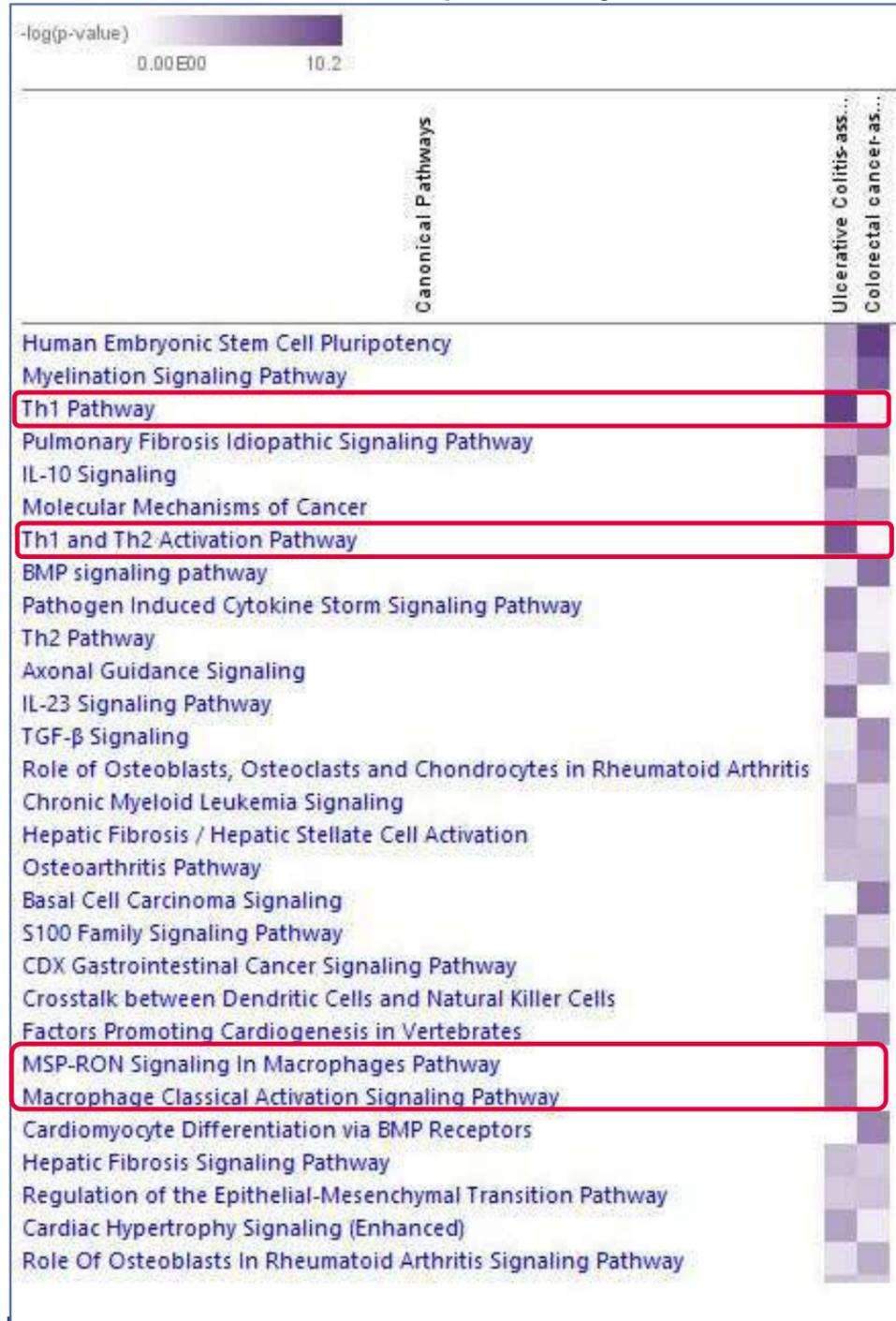


MSP-RON Signaling in Macrophages Pathway

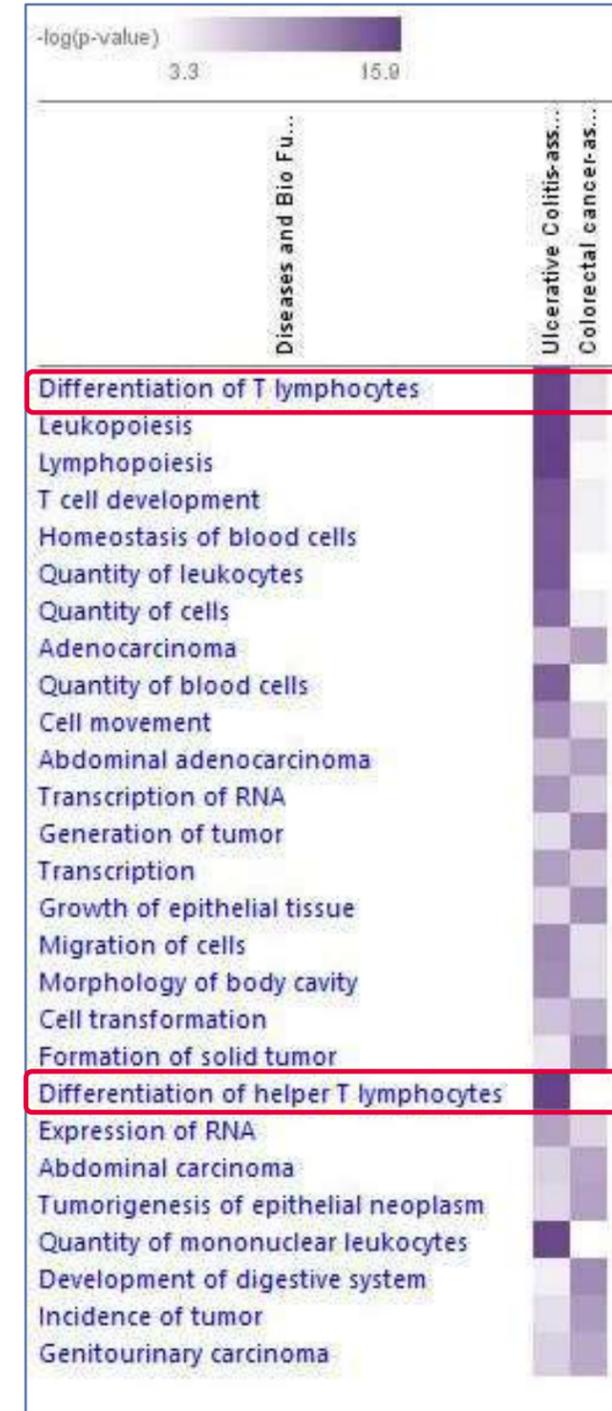




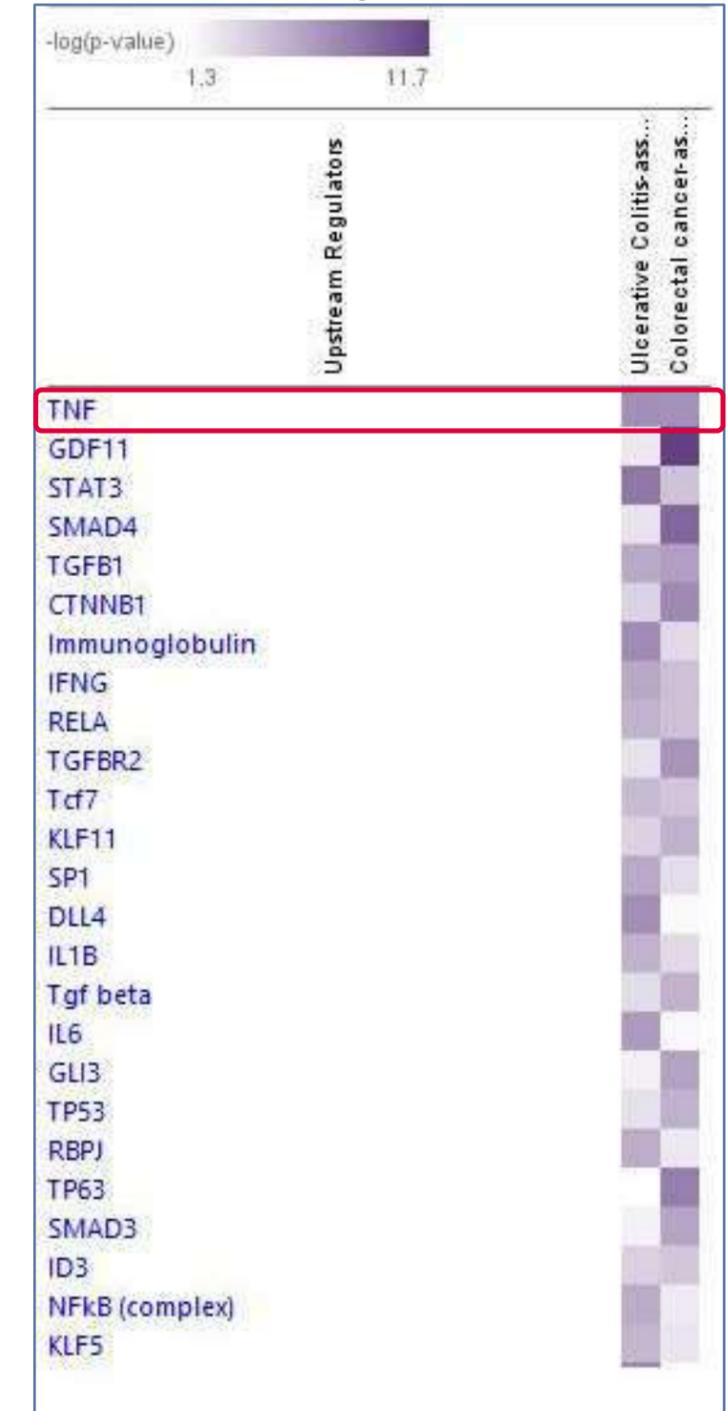
## Canonical pathways



## Disease and bio-functions



## Upstream Regulators



Compare
[-] [x]

Select Entities to compare and click Add

My Projects

- GWAS
  - Dataset Files
  - Analyses
    - Ulcerative Colitis-associations
    - Colorectal cancer-associations

Refresh

Add >>

<< Remove

Clear All

Calculate intersections

Click in the Venn diagram below to compare different sets.  
Keep the Ctrl key down to select multiple areas.

A Ulcerative Colitis-associations (Analysis)

B Colorectal cancer-associations (Analysis)

Entities Comparison Results (16)

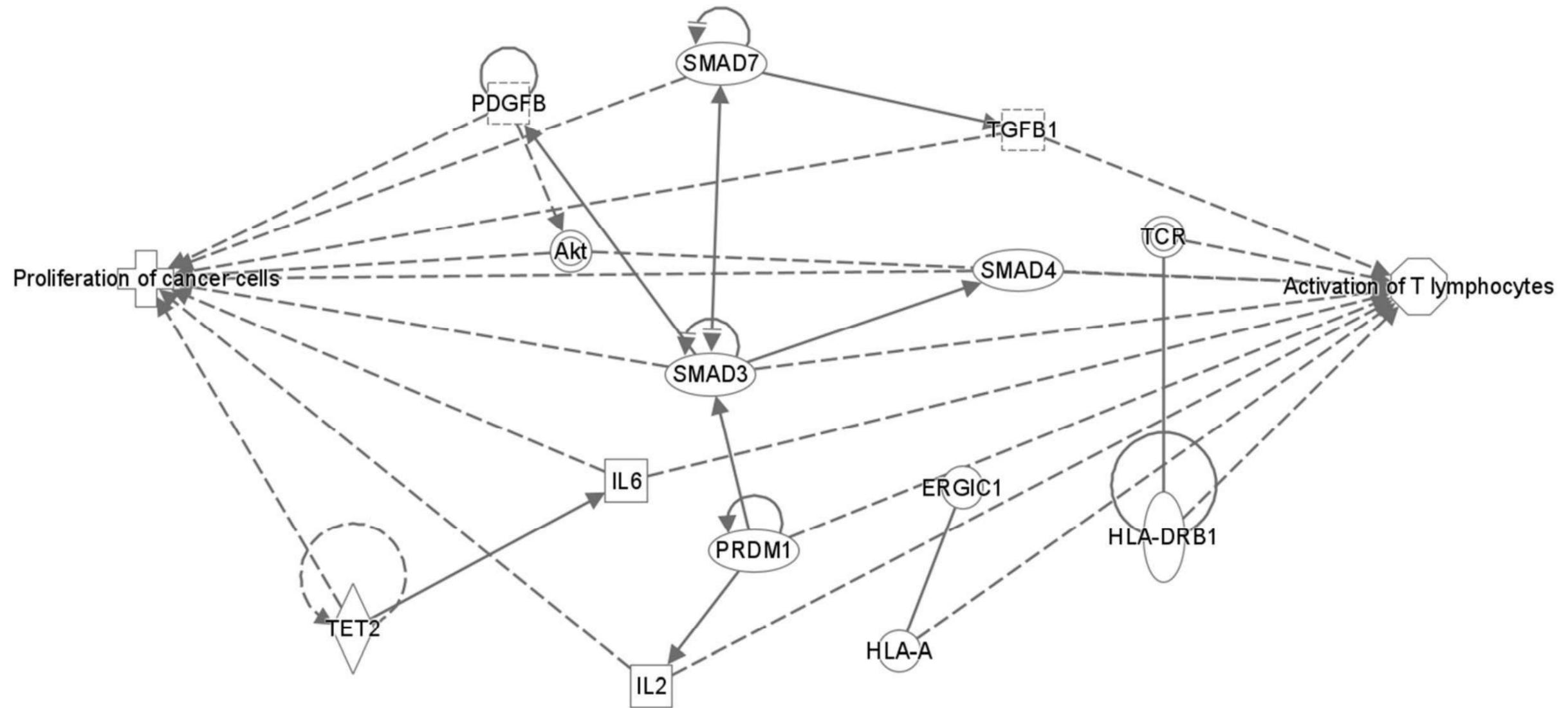
ERGIC1
GNA12
HLA-DRB1
LINC01271
LOC105373831
PDGFB
PITX1-AS1
PLCL1
PNKD
PRDM1
RPS21P8
SFMBT1
SMAD3
SMAD7
TET2
ZBTB40

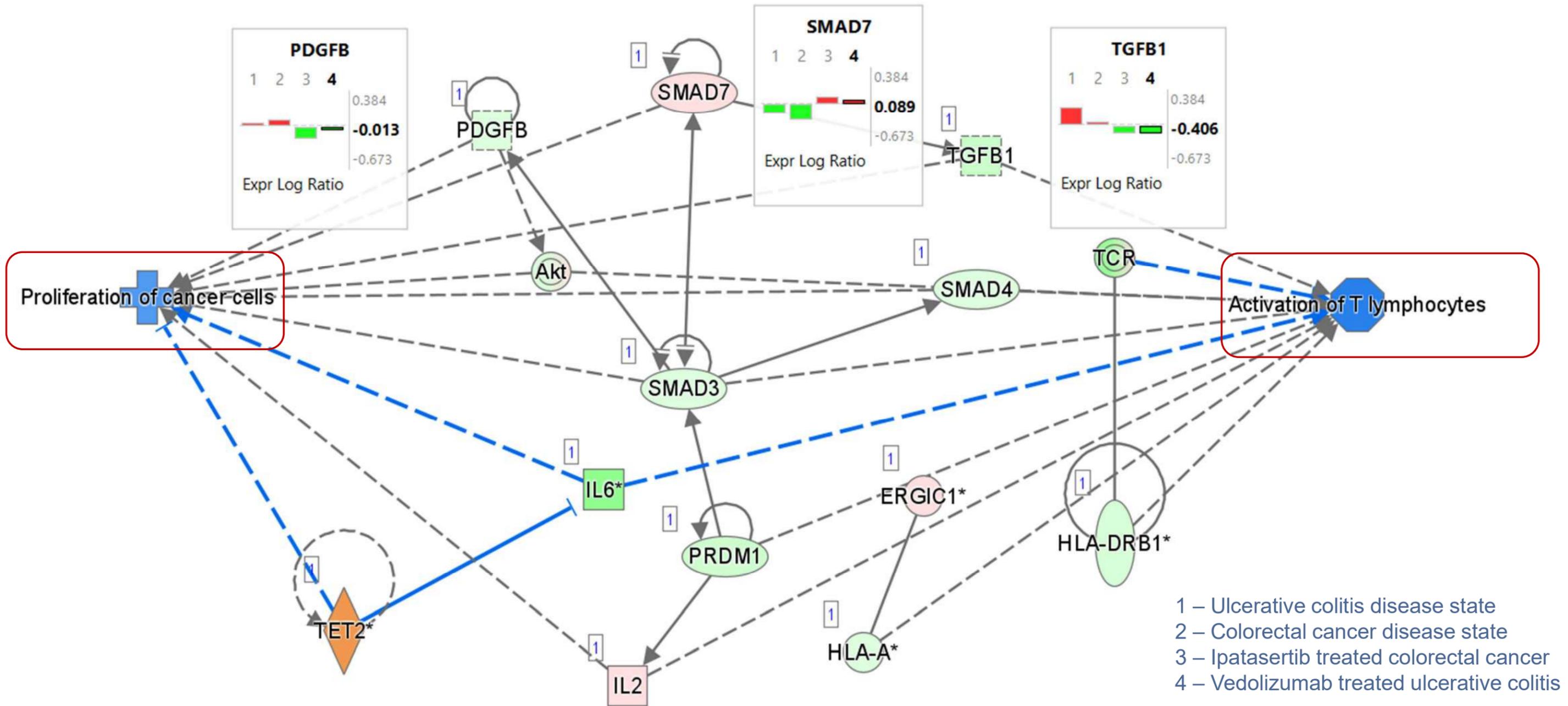
Show All

Add To My Pathway

Add To My List

Annotations

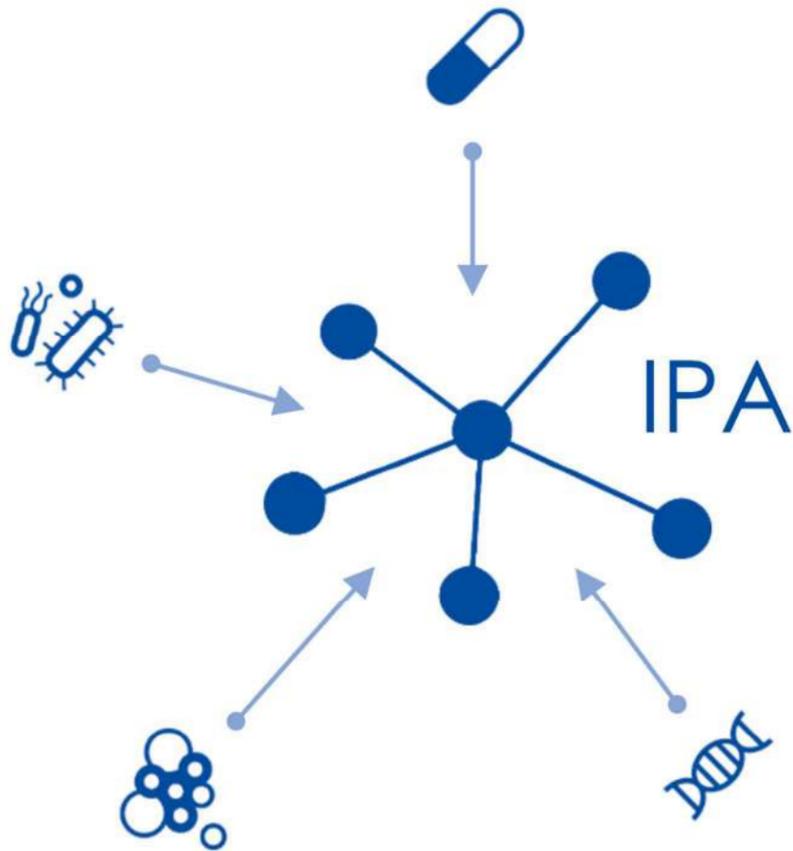




Updated weekly with the latest clinical trials and manuscripts

12+ million findings curated using controlled vocabulary

Decades of accumulated knowledge mapped in directional networks

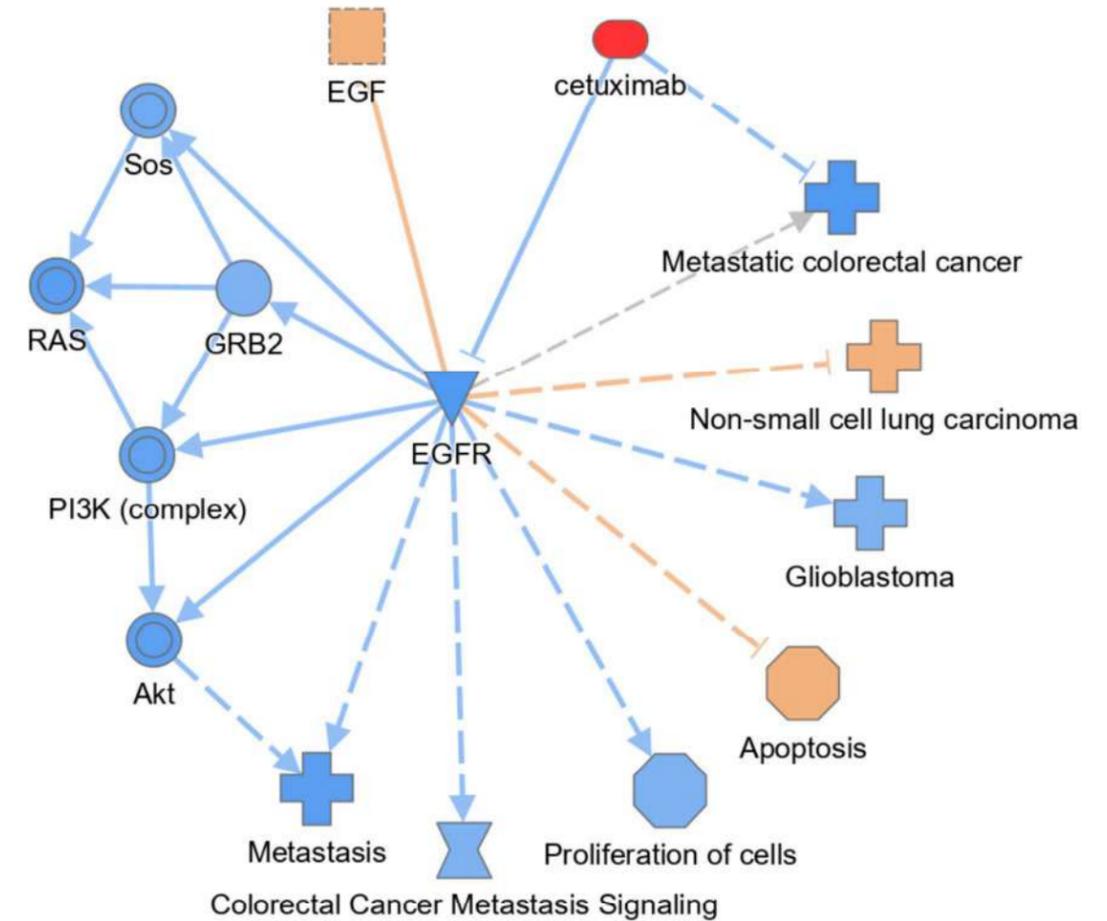


**EGF – EGFR**  
*[one of many]*

Type: **activation**  
 Direction: **directional**  
 Effect: **increases**  
 Directness: **direct**  
 Tissue: **epithelial cells**  
 Location: **plasma membrane**  
 Source: **PMID: 17909010**

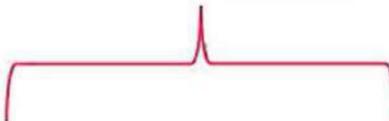
**cetuximab – EGFR**  
*[one of many]*

Type: **phosphorylation**  
 Direction: **directional**  
 Effect: **decreases**  
 Cell line: **CaR1 cells**  
 Organism: **human**  
 Experiment: **immunoblot**  
 Source: **PMID: 23213241**



	A
1	Risk loci
2	rs6012915
3	rs6095946
4	rs6091189
5	rs6014965
6	rs1741640
7	rs9983528
8	rs4616575
9	rs130651
10	rs5751474
11	rs34256596
12	rs9614460
13	rs9330814
14	rs2732875
15	rs174537
16	rs10751097
17	rs11236187
18	rs117042741
19	rs55864876
20	rs2155065
21	rs3087967
22	rs497916
23	rs10774214
24	rs3217810
25	rs3217874
26	rs10849432
27	rs10849434
28	rs77969132
29	rs2730985

Multiple ID columns

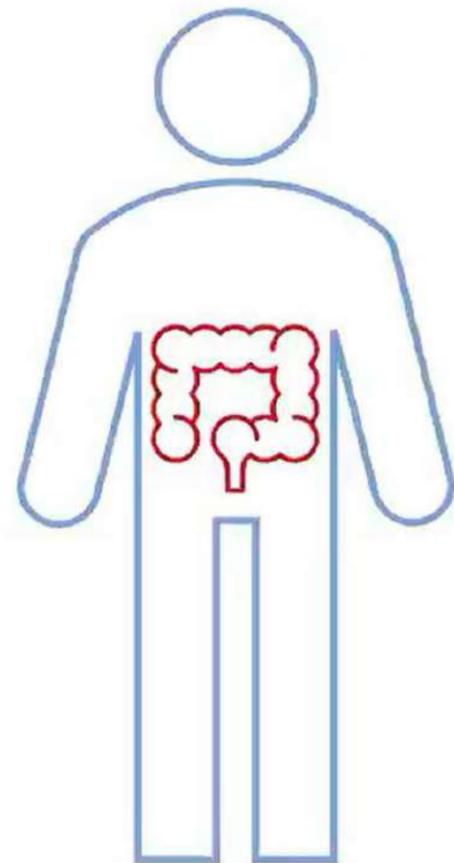


	A	B
1	Risk loci	Mapped gene
2	rs6012915	LINC01271
3	rs6095946	COX6CP2
4	rs6091189	RIPOR3
5	rs6014965	BMP7
6	rs1741640	LAMA5
7	rs9983528	PCNT
8	rs4616575	ZNRF3
9	rs130651	PDGFB
10	rs5751474	SCUBE1
11	rs34256596	LINC01639
12	rs9614460	SMC1B
13	rs9330814	WNT7B
14	rs2732875	SHROOM2
15	rs174537	MYRF
16	rs10751097	ANO1
17	rs11236187	POLD3
18	rs117042741	XRR1

## Accepted Files Formats

- .txt (tab delimited text files)
- .xls, .xlsx, .csv (excel files)
- .diff (cuffdiff files)

Analysis of genetic loci list identified in genome-wide association study (GWAS)



**Ulcerative Colitis**  
(PMID: 26192919)

	A	B	C	D	E	F
1	Variant	Mapped gene				
2	rs9847710	SFMBT1				
3	rs6087990	DNMT3B				
4	rs3740415	MFSD13A				
5	rs616597	NFKBIZ				
6	rs11150589	Y_RNA				
7	rs254560	C5orf66				
8	rs11641184	LITAF				
9	rs7657746	KIAA1109				
10	rs1405108	'-				
11	rs113010081	LINC02009				
12	rs12103	INTS11				
13	rs1517352	STAT4				
14	rs4976646	RGS14				
15	rs8005161	GPR65				
16	rs4656958	ITLN2				
17	rs4743820	LINC00484				
18	rs1077773	LINC02888				
19	rs907611	LSP1				
20	rs6062504	ZGPAT				
21	rs11229555	GLYAT				
22	rs913678	LINC01271				

Account: ipaXX@ingenuity.com

(XX= 01~30)

Password: **9-Str0ngP@&&IPA**

Expired until 10/20

*Download website:* <https://tinyurl.com/GGAIPA>

*Dataset:* <https://tinyurl.com/GGAdataset>

*Client: (IPA Login)*

Enter gene names/symbols/IDs or chemical/drug names here

Search

Advanced Search



Create New...

Core Analysis...

Comparison Analysis...

Biomarker Filter...

Biomarker Comparison Analysis...

MicroRNA Target Filter...

BioProfiler

IsoProfiler

My Pathway

Path Designer

Filter Dataset

Upload Dataset...

Advanced Search

Project...

Compare

Import Pathway

×

Refresh

Information

### Quick Start

News

Getting started

Exploring large public data resources through IPA

Exploring IPA knowledge

Analyzing mRNA or proteomics data

Analyzing microRNA data

Analyzing phosphoproteomics data

Analyzing genetic gain/loss data

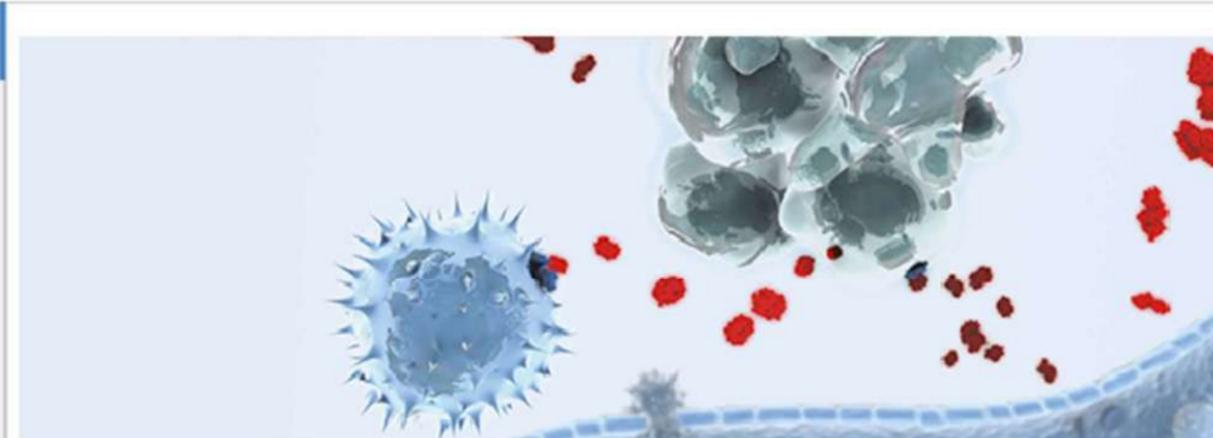
Analyzing metabolomics data

Case studies and Support webinars

Top help articles and FAQs

Contacting Support

Shortcuts



### News

- Starting in Jan 2024, analyses >10 years old will be eligible for [automatic deletion](#).
- Sign up for an [IPA training webinar!](#)
- Check out our NEW [1-minute IPA promo video](#).
- Check out the [single cell expression tutorial](#).
- Read and watch our [gene expression tutorial](#) with embedded videos.
- The IPA downtime for the upcoming **IPA 2023 Winter Release** has been scheduled for:
  - Pacific Standard Time: Friday, December 15, 5 p.m. through Sunday, December 17, 12 p.m. PST (Noon).
  - Central European Time: Saturday, December 16, 02:00 through Sunday, December 17, 21:00.
  - Japan Standard Time: Saturday, December 16, 10:00 through Monday, December 18, 05:00.
  - China CST (Beijing): Saturday, Saturday, December 16, 23:00 through Monday, December 18, 18:00.
- If you are new to IPA or taking a trial please see: [IPA Trial Resources](#).
- Pave your way to greatness using advanced pathway analysis: [Learn more](#).
- Read our [news](#) and sign up for our newsletter.
- Search Google Scholar for [publications that cite IPA](#).
- See how the IPA team is using Machine Learning in a [new paper](#).

Don't Show at Startup



Create New...

Search Advanced Search

Dataset Upload - Ulcerative Colitis-associations.csv



- 1. Select File Format: Flexible Format
- 2. Contains Column Header: Yes
- 3. Select Identifier Type: Please assign at least one column below as "ID", and assign the identifier type(s). Assign additional columns as ID to improve mapping coverage if desired.
- 4. Array platform used for experiments: Not specified/applicable
- 5. Use the dropdown menus to specify the column names that contain identifiers and observations. For observations, select the appropriate measurement value type.

Raw Data (502) Dataset Summary (335) Metadata

Edit Observation Names Infer Observations

ID/Observation Name	ID	Ignore
Measurement/Annotation	dbSNP	Gene Symbo...
1	Variant	
2	rs11230563	
3	rs11010067	
4	rs7556897	
5	rs11083840	
6	rs17736589	
7	rs17771967	
8	rs11054935	
9	rs11583043	
10	rs559928	
11	rs7011507	
12	rs11741861	
13	rs2777491	RTF1
14	rs13407913	ADCY3
15	rs28374715	CHP1
16	rs2488389	DENND1B
17	rs2930047	DAP
18	rs17293632	SMAD3
19	rs3116494	CD28
20	rs11064881	CIT

- Entrez Gene
- GenBank
- Gene Symbol - human (HUGO / HGNC / Entrez Gene)
- Gene Symbol - mouse (Entrez Gene)
- Gene Symbol - rat (Entrez Gene)
- GenPept
- GI Number
- Human Metabolome Database (HMDB)
- Illumina
- Inaenuitv

OK Cancel Clear

Save Cancel Help



- 1. Select File Format: Flexible Format
- 2. Contains Column Header: Yes
- 3. Select Identifier Type: Please assign at least one column below as "ID", and assign the identifier type(s). Assign additional columns as ID to improve mapping coverage if desired.
- 4. Array platform used for experiments: Not specified/applicable
- 5. Use the dropdown menus to specify the column names that contain identifiers and observations. For observations, select the appropriate measurement value type.

Raw Data (502) Dataset Summary (335) Metadata

ID/Observation Name	ID	ID	Ignore
	dbSNP	Gene Symbo...	
1	Variant	Mapped gene	----
2	rs11230563	CD6	----
3	rs11010067	LINC02635 - CUL2	----
4	rs7556897	SNRPGP8 - CCL20	----
5	rs11083840	CALM3 - PTGIR	----
6	rs17736589	CYTH1	----
7	rs17771967	KIR3DL2 - RNU6-222P	----
8	rs11054935	DUSP16	----
9	rs11583043	DPH5	----
10	rs559928	RPS6KA4 - LINC02723	----
11	rs7011507	UBE2V2 - IDI1P2	----
12	rs11741861	IRGM, ZNF300	----
13	rs2777491	RTF1	----
14	rs13407913	ADCY3	----
15	rs28374715	CHP1	----
16	rs2488389	DENND1B	----
17	rs2930047	DAP	----
18	rs17293632	SMAD3	----
19	rs3116494	CD28	----
20	rs11064881	CIT	----

Warning

Your dataset does not contain any metadata.  
Metadata is useful when you search for datasets or subsequent analyses.

Please click on the Metadata tab to add keywords that describe the data.

Click OK to continue to save the dataset without metadata.

OK Cancel

Save Cancel Help



Create New...

Search

Advanced Search

Dataset Upload - Ulcerative Colitis-associations.csv



1. Select File Format: Flexible Format
2. Contains Column Header:  Yes  No
3. Select Identifier Type: Please assign at least one column below as "ID", and assign the identifier type(s).  
Assign additional columns as ID to improve mapping coverage if desired.
4. Array platform used for experiments: Not specified/applicable Select relevant array platform as a reference set for data analysis.
5. Use the dropdown menus to specify the column names that contain identifiers and observations. For observations, select the appropriate measurement value type.

Raw Data (502) Dataset Summary (335) Metadata

Edit Observation Names Infer Observations

ID/Observation Name	ID	ID	Ignore
Measurement/Annotation	dbSNP	Gene Symbo...	
1	Variant	Mapped gene	---
2	rs11230563	CD6	---
3	rs11010067	LINC02635 - CUL2	---
4	rs7556897	SNRPGP8 - CCL20	---
5	rs11083840	CALM3 - PTGIR	---
6	rs17736589	CYTH1	---
7	rs17771967	KIR3DL2 - RNU6-222P	---
8	rs11054935	DUSP16	---
9	rs11583043	DPH5	---
10	rs559928	RPS6KA4 - LINC02723	---
11	rs7011507	UBE2V2 - IDI1P2	---
12	rs11741861	IRGM, ZNF300	---
13	rs2777491	RTF1	---
14	rs13407913	ADCY3	---
15	rs28374715	CHP1	---
16	rs2488389	DENND1B	---
17	rs2930047	DAP	---
18	rs17293632	SMAD3	---
19	rs3116494	CD28	---
20	rs11064881	CIT	---

Save Dataset

Choose Project: GWAS

Name: Ulcerative Colitis-associations

Notes:   
  
 (max 1600 chars)

Save

Cancel

Help



Create New...

Search

Advanced Search

Annotated Dataset: Ulcerative Colitis-associations2

Preview Dataset Ulcerative Colitis-associations2

Mapped IDs (335) Unmapped IDs (166) All IDs (501) Metadata

Add To My Pathway

Add To My List

Create Dataset

Customize Table



Symbol ACO2 - GNA12 (1/4)

ID	Flags	Symbol	Entrez Gene Name	Location	Type(s)	Drug(s)
rs727563	D	ACO2*	aconitase 2	Cytoplasm	enzyme	
rs727563	D	ACO2*	aconitase 2	Cytoplasm	enzyme	
rs2641348	D	ADAM30*	ADAM metalloproteinase domain 30	Plasma Membrane	peptidase	
rs2641348	D	ADAM30*	ADAM metalloproteinase domain 30	Plasma Membrane	peptidase	
rs13407913	D	ADCY3*	adenylate cyclase 3	Plasma Membrane	enzyme	
rs13407913	D	ADCY3*	adenylate cyclase 3	Plasma Membrane	enzyme	
rs13407913	D	ADCY3*	adenylate cyclase 3	Plasma Membrane	enzyme	
rs71624119	D	ANKRD55*	ankyrin repeat domain 55	Nucleus	transcription regulator	
rs10065637	D	ANKRD55*	ankyrin repeat domain 55	Nucleus	transcription regulator	
rs6738490	D	ATG16L1*	autophagy related 16 like 1	Cytoplasm	enzyme	
rs12994997	D	ATG16L1*	autophagy related 16 like 1	Cytoplasm	enzyme	
rs653178	D	ATXN2*	ataxin 2	Nucleus	other	
rs653178	D	ATXN2*	ataxin 2	Nucleus	other	
rs1847472	D	BACH2*	BTB domain and CNC homolog 2	Nucleus	transcription regulator	
rs1847472	D	BACH2*	BTB domain and CNC homolog 2	Nucleus	transcription regulator	
rs34592089	D	BANK1*	B cell scaffold protein with ankyrin repeats 1	Extracellular Space	other	
rs13126505	D	BANK1*	B cell scaffold protein with ankyrin repeats 1	Extracellular Space	other	
rs7657746	D	BLTP1*	bridge-like lipid transfer protein family me...	Other	other	
rs7657746	D	BLTP1*	bridge-like lipid transfer protein family me...	Other	other	
rs7657746	D	BLTP1*	bridge-like lipid transfer protein family me...	Other	other	
rs9836291		BSN	bassoon presynaptic cytomatrix protein	Plasma Membrane	other	
rs34856868	D	BTBD8*	BTB domain containing 8	Nucleus	other	
rs34856868	D	BTBD8*	BTB domain containing 8	Nucleus	other	
rs3853824	D	C17orf67*	chromosome 17 open reading frame 67	Other	other	
rs3853824	D	C17orf67*	chromosome 17 open reading frame 67	Other	other	

0 / 335

Flags:

"D" - Duplicates. Gene/Protein/Chemical identifiers marked with an asterisk indicate that multiple identifiers in the dataset file map to a single gene/chemical in the Global Molecular Network.

"O" - Override molecules. Gene/Protein/Chemical identifiers marked as "Override" are displayed with italic text.

"A" - Gene/Protein/Chemical ID marked as Absent. The gene/protein/chemical will not be used as a focus molecule or appear in networks unless you also explicitly override this flag with the Override column.

Edit Dataset Settings

Analyze/Filter Dataset

Close



Create New...

- Core Analysis...
- Comparison Analysis...
- Biomarker Filter...
- Biomarker Comparison Analysis...
- MicroRNA Target Filter...
- BioProfiler
- IsoProfiler
- My Pathway
- Path Designer
- Filter Dataset
- Upload Dataset...
- Advanced Search
- Project...
- Compare
- Import Pathway

Search

Advanced Search

All IDs (501) Metadata

List Create Dataset Customize Table

Symbol ACO2 - GNA12 (1/4)

Flags	Symbol	Entrez Gene Name	Location	Type(s)	Drug(s)
D	ACO2*	aconitase 2	Cytoplasm	enzyme	
D	ACO2*	aconitase 2	Cytoplasm	enzyme	
D	ADAM30*	ADAM metalloproteinase domain 30	Plasma Membrane	peptidase	
D	ADAM30*	ADAM metalloproteinase domain 30	Plasma Membrane	peptidase	
D	ADCY3*	adenylate cyclase 3	Plasma Membrane	enzyme	
D	ADCY3*	adenylate cyclase 3	Plasma Membrane	enzyme	
D	ADCY3*	adenylate cyclase 3	Plasma Membrane	enzyme	
D	ANKRD55*	ankyrin repeat domain 55	Nucleus	transcription regulator	
D	ANKRD55*	ankyrin repeat domain 55	Nucleus	transcription regulator	
D	ATG16L1*	autophagy related 16 like 1	Cytoplasm	enzyme	
D	ATG16L1*	autophagy related 16 like 1	Cytoplasm	enzyme	
D	ATXN2*	ataxin 2	Nucleus	other	
D	ATXN2*	ataxin 2	Nucleus	other	
D	BACH2*	BTB domain and CNC homolog 2	Nucleus	transcription regulator	
D	BACH2*	BTB domain and CNC homolog 2	Nucleus	transcription regulator	
D	BANK1*	B cell scaffold protein with ankyrin repeats 1	Extracellular Space	other	
D	BANK1*	B cell scaffold protein with ankyrin repeats 1	Extracellular Space	other	
D	BLTP1*	bridge-like lipid transfer protein family me...	Other	other	
D	BLTP1*	bridge-like lipid transfer protein family me...	Other	other	
D	BLTP1*	bridge-like lipid transfer protein family me...	Other	other	
D	BSN	bassoon presynaptic cytomatrix protein	Plasma Membrane	other	
D	BTBD8*	BTB domain containing 8	Nucleus	other	
D	BTBD8*	BTB domain containing 8	Nucleus	other	
D	C17orf67*	chromosome 17 open reading frame 67	Other	other	
D	C17orf67*	chromosome 17 open reading frame 67	Other	other	

0 / 335

Flags:  
 "D" - Duplicates. Gene/Protein/Chemical identifiers marked with an asterisk indicate that multiple identifiers in the dataset file map to a single gene/chemical in the Global Molecular Network.  
 "O" - Override molecules. Gene/Protein/Chemical identifiers marked as "Override" are displayed with italic text.  
 "A" - Gene/Protein/Chemical ID marked as Absent. The gene/protein/chemical will not be used as a focus molecule or appear in networks unless you also explicitly override this flag with the Override column.

Edit Dataset Settings Analyze/Filter Dataset Close



Create New...

Search

Advanced Search



Annotated Dataset: Ulcerative Colitis-associations2

Preview Dataset Ulcerative Colitis-associations2

Mapped IDs (335) Unmapped IDs (166) All IDs (501) Metadata

Add To My Pathway Add To My List Create Dataset Customize Table

Symbol ACO2 - GNA12 (1/4)

ID	Flags	Symbol	Type(s)	Drug(s)
rs727563	D	ACO2*	enzyme	
rs727563	D	ACO2*	enzyme	
rs2641348	D	ADAM30*	peptidase	
rs2641348	D	ADAM30*	peptidase	
rs13407913	D	ADCY3*	enzyme	
rs13407913	D	ADCY3*	enzyme	
rs13407913	D	ADCY3*	enzyme	
rs71624119	D	ANKRD55*	transcription regulator	
rs10065637	D	ANKRD55*	transcription regulator	
rs6738490	D	ATG16L1*	enzyme	
rs12994997	D	ATG16L1*	enzyme	
rs653178	D	ATXN2*	other	
rs653178	D	ATXN2*	other	
rs1847472	D	BACH2*	transcription regulator	
rs1847472	D	BACH2*	transcription regulator	
rs34592089	D	BANK1*	other	
rs13126505	D	BANK1*	other	
rs7657746	D	BLTP1*	other	
rs7657746	D	BLTP1*	other	
rs7657746	D	BLTP1*	other	
rs9836291		BSN	other	
rs34856868	D	BTBD8*	other	
rs34856868	D	BTBD8*	other	
rs3853824	D	C17orf67*	other	
rs3853824	D	C17orf67*	other	

Create Core Analysis

Upload

- My Projects
  - GWAS
    - Ulcerative Colitis-associations2
    - myList
    - colon cancer-association
    - Ulcerative Colitis-associations**
- Shared Projects
  - Projects Shared with Others

Next

0 / 335

Flags:

"D" - Duplicates. Gene/Protein/Chemical identifiers marked with an asterisk indicate that multiple identifiers in the dataset file map to a single gene/chemical in the Global Molecular Network.

"O" - Override molecules. Gene/Protein/Chemical identifiers marked as "Override" are displayed with italic text.

"A" - Gene/Protein/Chemical ID marked as Absent. The gene/protein/chemical will not be used as a focus molecule or appear in networks unless you also explicitly override this flag with the Override column.

Edit Dataset Settings Analyze/Filter Dataset Close





Create New...

Search

Advanced Search

Create Expression Analysis - [analysis : Ulcerative Colitis-associations]

General Settings

Population of genes to consider for p-value calculations:

Reference Set Ingenuity Knowledge Base (Genes Only)

Relationships to consider:

Affects networks and upstream regulator analysis

Direct and Indirect Relationships

Direct Relationships

Optional Analyses:

My Project

My Pathways

My Lists

Analysis Filter Summary

Consider only relationships where confidence = Experimentally Observed

Advanced

Save As Default

Recalculate

167 analysis-ready molecules across observations

Preview Dataset Ulcerative Colitis-associations

Analysis-Ready (167) Mapped IDs (168) Unmapped IDs (85) All IDs (253) Metadata

Add To My Pathway

Add To My List

Create Dataset

Customize Table



Symbol ACO2 - MYRF (1/2)

ID	Flags	Symbol	Entrez Gene Name	Location	Type(s)	Drug(s)
rs727563		<b>ACO2</b>	aconitase 2	Cytoplasm	enzyme	
rs2641348		<b>ADAM30</b>	ADAM metalloproteinase domain 30	Plasma Membrane	peptidase	
rs13407913		<b>ADCY3</b>	adenylate cyclase 3	Plasma Membrane	enzyme	
rs10065637		<b>ANKRD55</b>	ankyrin repeat domain 55	Nucleus	transcription regulator	
rs12994997		<b>ATG16L1</b>	autophagy related 16 like 1	Cytoplasm	enzyme	
rs653178		<b>ATXN2</b>	ataxin 2	Nucleus	other	
rs1847472		<b>BACH2</b>	BTB domain and CNC homolog 2	Nucleus	transcription regulator	

0 / 167

Flags: "Bold" Focus molecules. Gene/Protein/Chemical identifiers that meet the user-defined cutoff and map to the Global Molecular Network are displayed with bold text.

Run Analysis

Cancel

Expression Analysis - Ulcerative Colitis-associations - 2023-10-19 11:07 上午

Summary Graphical Summary Pathways Upstream Analysis Diseases & Functions Regulator Effects Networks Lists Analysis Match Molecules

The analysis has started.  
Tabs will be enabled as soon as they complete.

Tab	Estimated completion
Molecules	Done
Canonical Pathways	Done
My Pathways	Done
ML Disease Pathways	Done
My Lists	Done
Tox Lists	Done
Diseases & Functions	< 30 minutes
Upstream Regulators	< 30 minutes
Regulator Effects	< 30 minutes
Graphical Summary	< 30 minutes
Causal Networks	< 30 minutes
Networks	< 35 minutes



Expression Analysis - Ulcerative Colitis-associations - 2023-10-18 05:19 下午

Summary Graphical Summary Pathways Upstream Analysis Diseases & Functions Regulator Effects Networks Lists Analysis Match Molecules

Export:

> Experiment Metadata

> Analysis Settings

> Top Canonical Pathways

Name	p-value	Overlap
Th1 Pathway	2.51E-10	8.9 % 11/124
Th1 and Th2 Activation Pathway	7.03E-10	6.9 % 12/174
Natural Killer Cell Signaling	3.44E-09	6.0 % 12/200
Role of JAK1, JAK2 and TYK2 in Interferon Signaling	9.11E-09	23.1 % 6/26
IL-23 Signaling Pathway	1.52E-08	14.6 % 7/48

> Top Upstream Regulators

> Upstream Regulators

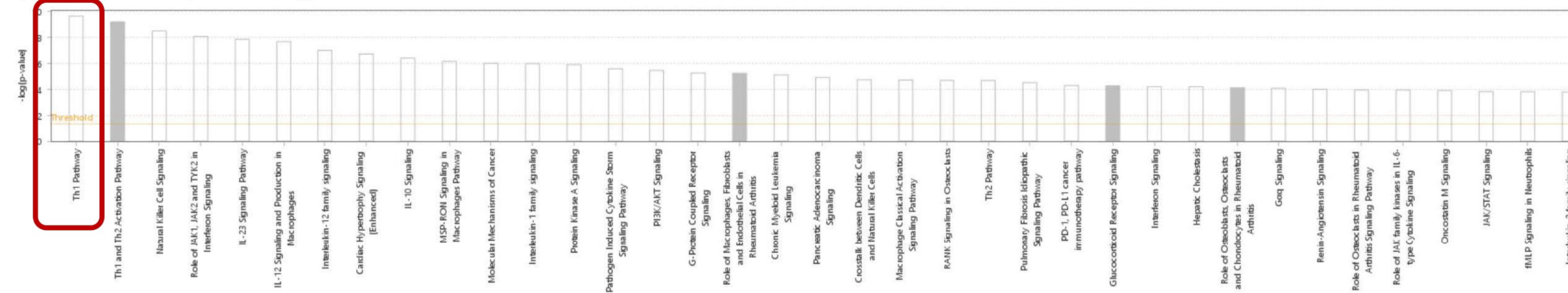
Name	p-value	Predicted Activation
lipopolysaccharide	5.33E-16	
IL12 (complex)	1.28E-12	
CD28	2.69E-12	
TCR	7.23E-12	
prostaglandin E2	2.93E-10	

> Causal Network

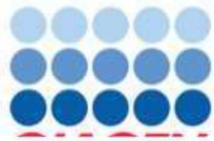
Name	p-value	Predicted Activation
IL23A	3.70E-13	
CDYL2	2.41E-12	
CRLF1	3.28E-12	
CD3-TCR	3.72E-12	
APOC1	3.84E-12	

> Top Diseases and Bio Functions

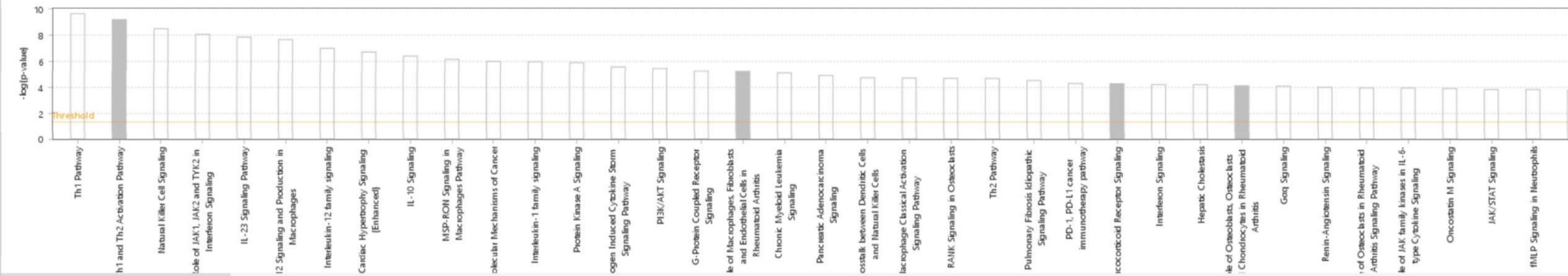
> Diseases and Disorders



Click on a data point in the chart above to see which molecules are associated with that pathway



positive z-score z-score = 0 negative z-score no activity pattern available



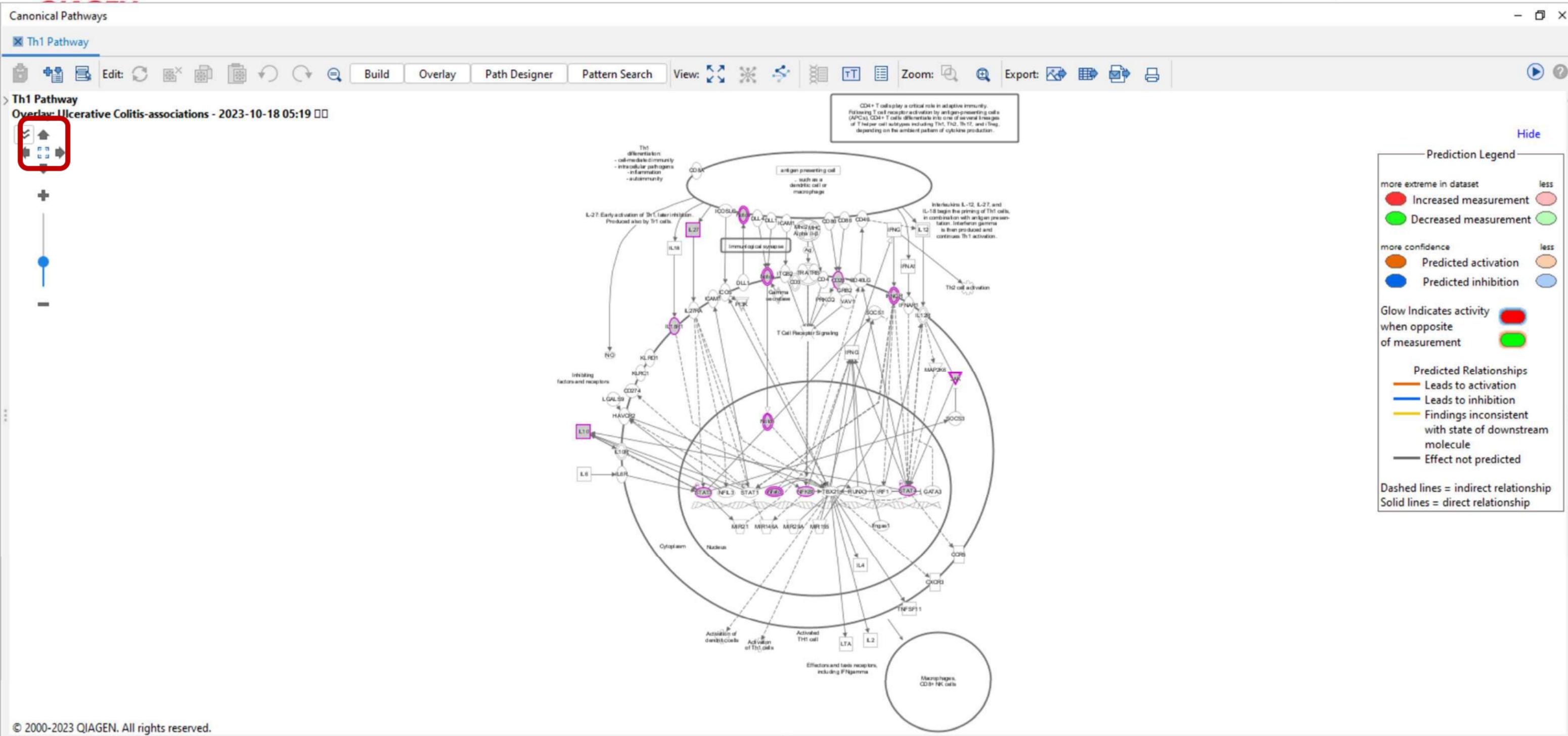
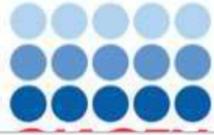
11 molecule(s) associated with Th1 Pathway [Ratio: 11/124 (0.089)] [z-score: NaN] [p-value: 2.51E-10]

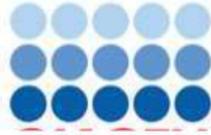
Activity Plot View Report Open Pathway

Add To My Pathway Add To My List Create Dataset Customize Table

Symbol	Entrez Gene Name	Identifier	Expected	Location	Type(s)	Biomarker Application(s)	Drug(s)
CD28	CD28 molecule	rs3116494	↑ Up	Plasma Membrane	transmembrane receptor	efficacy	FPT155, lulizumab pegol, REGN... all 8
IFNGR2	interferon gamma receptor 2	rs2284553	↑ Up	Plasma Membrane	transmembrane receptor		interferon gamma-1b all 1
IL10	interleukin 10	rs3024493	↑ Up	Extracellular Space	cytokine	diagnosis, disease progression, .....all 7	SCH708980 all 1
IL27	interleukin 27	rs26528	↑ Up	Extracellular Space	cytokine		SRF388 all 1
IL18R1	interleukin 18 receptor 1	rs1420098	↑ Up	Plasma Membrane	transmembrane receptor		
NFATC1	nuclear factor of activated T cells 1	NFATC1	↑ Up	Nucleus	transcription regulator		
NFKB1	nuclear factor kappa B subunit 1	rs3774937	↑ Up	Nucleus	transcription regulator		bortezomib/dexamethasone/th... all 7
NOTCH1	notch receptor 1	rs13300218	↑ Up	Plasma Membrane	transcription regulator	diagnosis, efficacy	OMP-52M51 all 1

Selected/Total molecules: 0 / 11





Upstream Regulator	Molecule Type	n-value of overlap	Target Molecules in Dataset	Mechanistic Network
lipopolysaccharide	chemical drug		BACH2, BANK1, CD28, CDC37, CPEB4, CREM, DUSP16, ER... ..all 52	73 (20)
IL12 (complex)	complex		CD226, CD28, IFIH1, IFNGR2, IL10, IL18R1, IL18RAP, IL27, ...all 15	64 (20)
CD28	transmembrane receptor		CD226, CD28, CPEB4, CREM, IL10, IL18R1, IL23R, IL27, IL2... ..all 18	47 (16)
TCR	complex		BACH2, CD28, CD6, F5, IFIH1, IL10, IL18R1, IL2RA, MAP3K8, ...all 18	65 (22)
prostaglandin E2	chemical - endogenous mammalian		CD226, CREM, CYTH1, FOSL2, IL10, IL23R, IL27, IL2RA, M... ..all 17	64 (21)
Immunoglobulin	complex		ACO2, ADCY3, BACH2, CD226, CD28, ERGIC1, FADS2, FC... ..all 27	58 (19)
Notch	group		CARD9, FCGR2A, IL10, LRRK2, LTBR, NFATC1, NOTCH1, P... ..all 11	57 (18)
TNF	cytokine		ANKRD55, CD28, CREM, DUSP16, FOSL2, GATD3/LOC102... ..all 37	63 (20)
LMO2	transcription regulator		CARD9, EPHB4, IFIH1, IL2RA, LSP1, LTBR, MAP3K8, NOTC... ..all 14	
IFNG	cytokine		BACH2, CREM, ERAP2, FCGR2A, IFIH1, IFNGR2, IL10, IL18... ..all 30	67 (23)
IL21	cytokine		BACH2, CD28, IL10, IL18R1, IL18RAP, IL23R, IL2RA, PRDM1, ...all 11	54 (17)
tretinoin	chemical drug		BACH2, CD6, CREM, DAP3, DNMT3B, FOSL2, GNA12, IFIH1, ...all 32	75 (22)
tetradecanoylphorbol acetate	chemical drug		ADCY3, CD226, CD28, CREM, FOSL2, GPR35, GPR65, IFN... ..all 26	58 (18)
STAT3	transcription regulator		BACH2, CD226, DNMT3B, GPR65, IFIH1, IL10, IL18R1, IL23R, ...all 21	57 (19)
IgG	complex		BANK1, FCGR2A, FOSL2, IFIH1, IFNGR2, IL10, IL2RA, ITGAL, ...all 14	67 (20)
REL	transcription regulator		BACH2, CREM, FADS2, IL27, IL2RA, MAP3K8, NFATC1, NF... ..all 12	56 (14)
IL2	cytokine	2.07E-08	ADCY3, CARD9, CD28, CPEB4, DAP, FADS2, FOSL2, IFNGR2, ...all 20	57 (17)
NFkB (complex)	complex	4.25E-08	CREM, ERAP2, HGFAC, IL10, IL27, IL2RA, ITGAL, LITAF, LS... ..all 18	62 (19)
TNFRSF21	transmembrane receptor	4.56E-08	CD28, IL10, IL2RA, NFATC1, STAT3 ...all 5	46 (11)
LDB1	transcription regulator	5.28E-08	CARD9, EPHB4, IFIH1, IL2RA, ITGAL, LSP1, LTBR, MAP3K8, ...all 13	
GNAS	enzyme	9.43E-08	BACH2, CREM, FOSL2, HDAC7, IL2RA, LITAF, PLA2G4A, P... ..all 10	51 (8)
STAT4	transcription regulator	9.60E-08	DAP, IFIH1, IL10, IL18R1, IL18RAP, IL2RA, MAP3K8, PRDM1, ...all 11	56 (18)
IL23A	cytokine	1.19E-07	IL10, IL18R1, IL23R, NFATC1, RORC, STAT3 ...all 6	49 (16)
FOXP3	transcription regulator	1.28E-07	CD226, FOSL2, IFNGR2, IL10, IL18RAP, IL2RA, NFKB1, PRD... ..all 10	58 (21)
RELA	transcription regulator	1.91E-07	BACH2, CD28, DNMT3B, ERAP2, IFNGR2, IL10, IL27, IL2RA, ...all 15	58 (21)
resiquimod	chemical drug	2.46E-07	CD226, FCGR2A, IL10, IL18RAP, IL27, IL2RA, NFKB1, NFKBIZ, ...all 14	62 (21)
IRF4	transcription regulator	2.62E-07	BACH2, CD28, IL10, IL18RAP, IL23R, IL2RA, ITGAL, NFKB1, ...all 10	54 (20)
IL18	cytokine	2.62E-07	CD226, FOSL2, IL10, IL18R1, IL18RAP, IL2RA, ITGAL, NFKBIZ, ...all 10	59 (21)
ZBTB10	transcription regulator	3.44E-07	CARD9, IFIH1, IL10, IL27, IL2RA, LRRK2, PTPN22, RELA, TNF... ..all 9	
DLL4	other	3.56E-07	EPHB4, IL10, NFKB1, NOTCH1, RELA, RORC ...all 6	61 (15)
IL1B	cytokine	3.58E-07	CREM, FADS2, IFNGR2, IL10, IL18R1, IL18RAP, IL23R, IL27, ...all 23	61 (18)
IL4	cytokine	3.61E-07	ANKRD55, CD6, CYTH1, EPHB4, FCGR2A, IFIH1, IFNGR2, I... ..all 24	64 (20)
IL10	cytokine	6.47E-07	CD28, CIT, FCGR2A, IFNGR2, IL10, IL18R1, IL27, ITGAL, M... ..all 14	65 (20)
IRAK1BP1	other	7.25E-07	IL10, NFKB1, RELA ...all 3	49 (12)

Molecule Type

Unfiltered

Genes, RNAs, and Proteins

Drugs and Chemicals

Select from list below

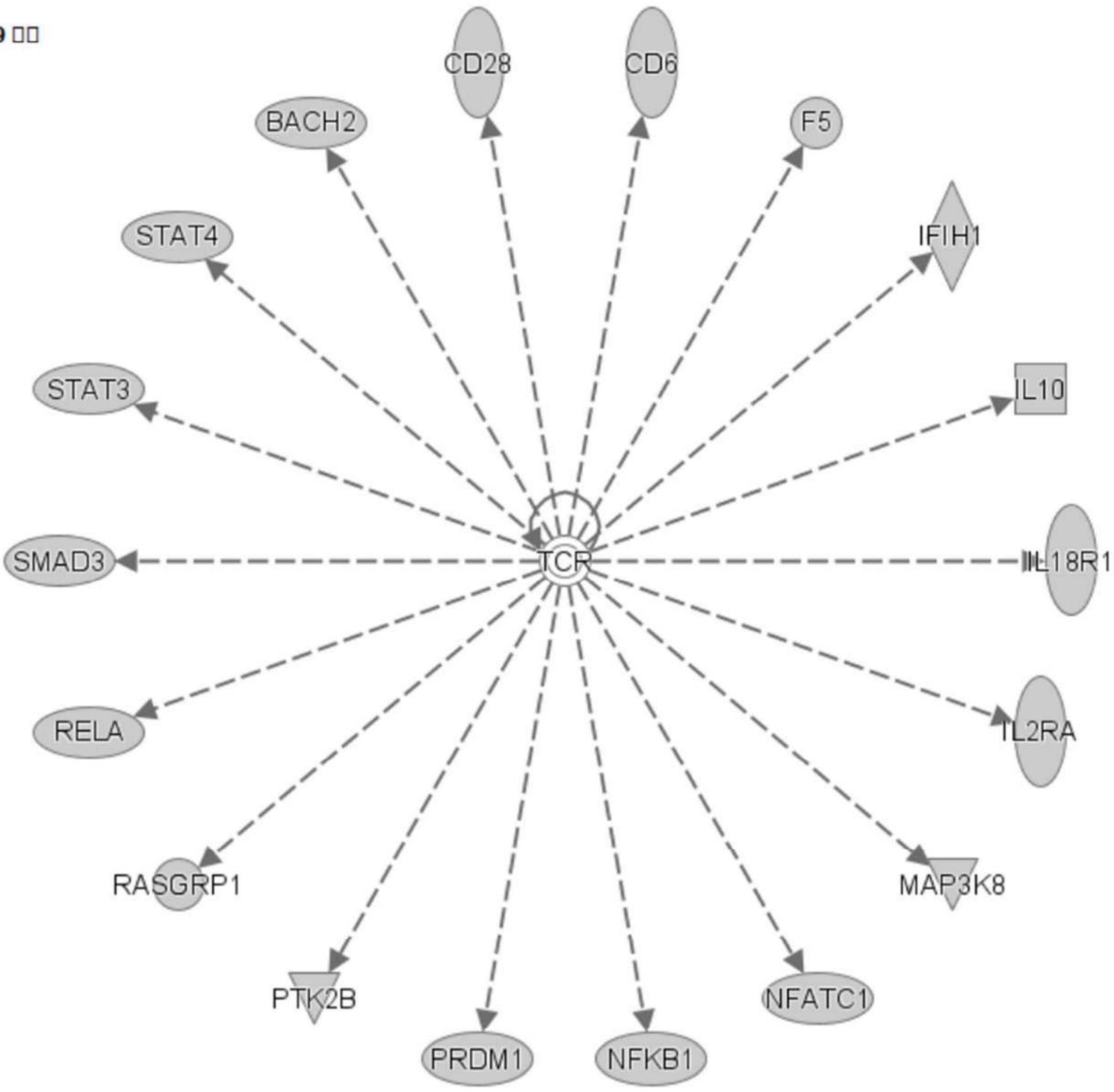
Select all

- biologic drug
- canonical pathway
- chemical - endogenous mammalian
- chemical - endogenous non-mammalian
- chemical - kinase inhibitor
- chemical - other

Apply Cancel

Upstream Regulator	Molecule Type	p-value of overlap	Target Molecules in Dataset	Mechanistic Network
lipopolysaccharide	chemical drug	5.33E-16	BACH2, BANK1, CD28, CDC37, CPEB4, CREM, DUSP16, ER... all 52	73 (20)
IL12 (complex)	complex	1.28E-12	CD226, CD28, IFIH1, IFNGR2, IL10, IL18R1, IL18RAP, IL27, ... all 15	64 (20)
CD28	transmembrane receptor	2.69E-12	CD226, CD28, CPEB4, CREM, IL10, IL18R1, IL23R, IL27, IL2... all 18	47 (16)
<b>TCR</b>	<b>complex</b>	<b>7.23E-12</b>	<b>BACH2, CD28, CD6, F5, IFIH1, IL10, IL18R1, IL2RA, MAP3K8, ... all 18</b>	<b>65 (22)</b>
prostaglandin E2	chemical - endogenous mammalian	2.93E-10	CD226, CREM, CYTH1, FOSL2, IL10, IL23R, IL27, IL2RA, M... all 17	64 (21)
Immunoglobulin	complex	3.61E-10	ACO2, ADCY3, BACH2, CD226, CD28, ERGIC1, FADS2, FC... all 27	58 (19)
Notch	group	9.33E-10	CARD9, FCGR2A, IL10, LRRK2, LTBR, NFATC1, NOTCH1, P... all 11	57 (18)
TNF	cytokine	9.61E-10	ANKRD55, CD28, CREM, DUSP16, FOSL2, GATD3/LOC102... all 37	63 (20)
LMO2	transcription regulator	1.81E-09	CARD9, EPHB4, IFIH1, IL2RA, LSP1, LTBR, MAP3K8, NOTC... all 14	
IFNG	cytokine	7.25E-09	BACH2, CREM, ERAP2, FCGR2A, IFIH1, IFNGR2, IL10, IL18... all 30	67 (23)
IL21	cytokine	7.37E-09	BACH2, CD28, IL10, IL18R1, IL18RAP, IL23R, IL2RA, PRDM1, ... all 11	54 (17)
tretinoin	chemical drug	9.37E-09	BACH2, CD6, CREM, DAP3, DNMT3B, FOSL2, GNA12, IFIH1, ... all 32	75 (22)
tetradecanoylphorbol acetate	chemical drug	9.90E-09	ADCY3, CD226, CD28, CREM, FOSL2, GPR35, GPR65, IFN... all 26	58 (18)
STAT3	transcription regulator	1.02E-08	BACH2, CD226, DNMT3B, GPR65, IFIH1, IL10, IL18R1, IL23R, ... all 21	57 (19)
IgG	complex	1.63E-08	BANK1, FCGR2A, FOSL2, IFIH1, IFNGR2, IL10, IL2RA, ITGAL, ... all 14	67 (20)
REL	transcription regulator	1.97E-08	BACH2, CREM, FADS2, IL27, IL2RA, MAP3K8, NFATC1, NF... all 12	56 (14)
IL2	cytokine	2.07E-08	ADCY3, CARD9, CD28, CPEB4, DAP, FADS2, FOSL2, IFNGR2, ... all 20	57 (17)
NFkB (complex)	complex	4.25E-08	CREM, ERAP2, HGFAC, IL10, IL27, IL2RA, ITGAL, LITAF, LS... all 18	62 (19)
TNFRSF21	transmembrane receptor	4.56E-08	CD28, IL10, IL2RA, NFATC1, STAT3	all 5 46 (11)
LDB1	transcription regulator	5.28E-08	CARD9, EPHB4, IFIH1, IL2RA, ITGAL, LSP1, LTBR, MAP3K8, ... all 13	
GNAS	enzyme	9.43E-08	BACH2, CREM, FOSL2, HDAC7, IL2RA, LITAF, PLA2G4A, P... all 10	51 (8)
STAT4	transcription regulator	9.60E-08	DAP, IFIH1, IL10, IL18R1, IL18RAP, IL2RA, MAP3K8, PRDM1, ... all 11	56 (18)
IL23A	cytokine	1.19E-07	IL10, IL18R1, IL23R, NFATC1, RORC, STAT3	all 6 49 (16)
FOXP3	transcription regulator	1.28E-07	CD226, FOSL2, IFNGR2, IL10, IL18RAP, IL2RA, NFKB1, PRD... all 10	58 (21)

[Navigation icons: pan, zoom, fit, etc.]



**Prediction Legend**

more extreme in dataset      less  
 ● Increased measurement      ●  
 ● Decreased measurement      ●

more confidence      less  
 ● Predicted activation      ●  
 ● Predicted inhibition      ●

Glow Indicates activity when opposite of measurement  
 ●  
 ●

**Predicted Relationships**  
 — Leads to activation  
 — Leads to inhibition  
 — Findings inconsistent with state of downstream molecule  
 — Effect not predicted

Dashed lines = indirect relationship  
 Solid lines = direct relationship





Symbol	Entrez Gene Name	Identifier	Location	Type(s)	Biomarker Application(s)	Drug(s)
ACO2	aconitase 2	rs727563	Cytoplasm	enzyme		
ADAM30	ADAM metallopeptidase domain 30	rs2641348	Plasma Membrane	peptidase		
ADCY3	adenylate cyclase 3	rs13407913	Plasma Membrane	enzyme		
ANKRD55	ankyrin repeat domain 55	rs10065637	Nucleus	transcription regulator		
ATG16L1	autophagy related 16 like 1	rs12994997	Cytoplasm	enzyme		
ATXN2	ataxin 2	rs653178	Nucleus	other		
BACH2	BTB domain and CNC homolog 2	rs1847472	Nucleus	transcription regulator		
BANK1	B cell scaffold protein with ankyrin repeats 1	rs13126505	Extracellular Space	other		
BLTP1	bridge-like lipid transfer protein family mem...	rs7657746	Other	other		
BSN	bassoon presynaptic cytomatrix protein	rs9836291	Plasma Membrane	other		
BTBD8	BTB domain containing 8	rs34856868	Nucleus	other		
C17orf67	chromosome 17 open reading frame 67	rs3853824	Other	other		
C3orf84	chromosome 3 open reading frame 84	rs4541435	Other	other		
CARD9	caspase recruitment domain family member 9	rs10781499	Cytoplasm	other	diagnosis	
CCDC26	CCDC26 long non-coding RNA	rs13277237	Other	other		
CD226	CD226 molecule	rs17207042	Plasma Membrane	other		LY3435151
CD28	CD28 molecule	rs3116494	Plasma Membrane	transmembrane receptor	efficacy	FPT155, REGN5668, REGN5678,
CD6	CD6 molecule	CD6	Plasma Membrane	transmembrane receptor		
CDC37	cell division cycle 37, HSP90 cochaperone	rs11879191	Cytoplasm	other		
CDC42SE2	CDC42 small effector 2	rs11743851	Plasma Membrane	other		
CDKAL1	CDK5 regulatory subunit associated protein...	rs2328546	Cytoplasm	enzyme		
CELSR3	cadherin EGF LAG seven-pass G-type recept...	rs9868809	Plasma Membrane	G-protein coupled receptor		
CHP1	calcineurin like EF-hand protein 1	rs28374715	Nucleus	transcription regulator		voclosporin
CIT	citron rho-interacting serine/threonine kina...	rs11064881	Cytoplasm	kinase		
CPEB4	cytoplasmic polyadenylation element bindin...	rs56163845	Plasma Membrane	other		
CREM	cAMP responsive element modulator	rs34779708	Nucleus	transcription regulator		
CYTH1	cytohesin 1	rs17736589	Cytoplasm	other	unspecified application	
DAP	death associated protein	rs2930047	Cytoplasm	transcription regulator		
DAP3	death associated protein 3	rs490608	Cytoplasm	other		
DELEC1	deleted in esophageal cancer 1	DELEC1	Other	other		
DENND1B	DENN domain containing 1B	rs2488389	Other	other		
DNMT3B	DNA methyltransferase 3 beta	rs6058869	Nucleus	enzyme		
DOCK7	dedicator of cytokinesis 7	rs1748195	Plasma Membrane	other		
DPH5	diphthamide biosynthesis 5	rs11583043	Cytoplasm	enzyme		
DUSP16	dual specificity phosphatase 16	rs11054935	Nucleus	phosphatase		

Selected/Total molecules: 0 / 167

Secondary **colon cancer** in patients with **ulcerative colitis**: a systematic review and meta-analysis.

Zhang L, Gan H.

J Gastrointest Oncol. 2021 Dec;12(6):2882-2890. doi: 10.21037/jgo-21-800.

A young woman who developed ascending **colon cancer** 2 years after the onset of **ulcerative colitis**.

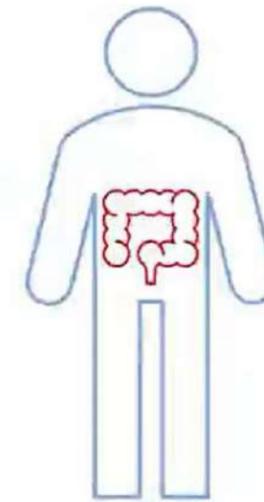
Matsuda K, Hashiguchi Y, Asako K, Okada Y, Ohno K, Tsukamoto M, Fukushima Y, Shimada R, Ozawa T, Hayama T, Nozawa K, Fukagawa T, Kikuchi Y, Sasajima Y.

Clin J Gastroenterol. 2020 Dec; 13(6):1189-1195

Risk of **Colon Cancer** and Recommended Surveillance Strategies in Patients with **Ulcerative Colitis**.

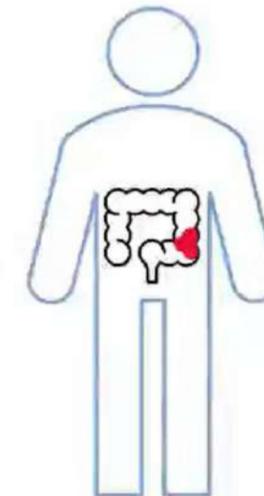
Rabbenou W, Ullman TA.

Gastroenterol Clin North Am. 2020 Dec;49(4):791-807



**Ulcerative Colitis GWAS**

(PMID: 26192919)



**Colorectal Cancer GWAS**

(PMID: 36539618)



Create New...

Search

Advanced Search

- Core Analysis...
- Comparison Analysis...
- Biomarker Filter...
- Biomarker Comparison Analysis...
- MicroRNA Target Filter...
- BioProfiler
- IsoProfiler
- My Pathway
- Path Designer
- Filter Dataset
- Upload Dataset...
- Advanced Search
- Project...
- Compare
- Import Pathway

- My Lists
  - Human Genes Chromosomal Location
  - QIAGEN Coronavirus Networks
  - Ingenuity KEGG gene lists
  - Tissue Expression
  - Example Analyses
  - Training
- Shared Projects

associations - 2023-10-19 11:07

association - 2023-10-18 05:52

associations - 2023-10-18 05:19

es

ults

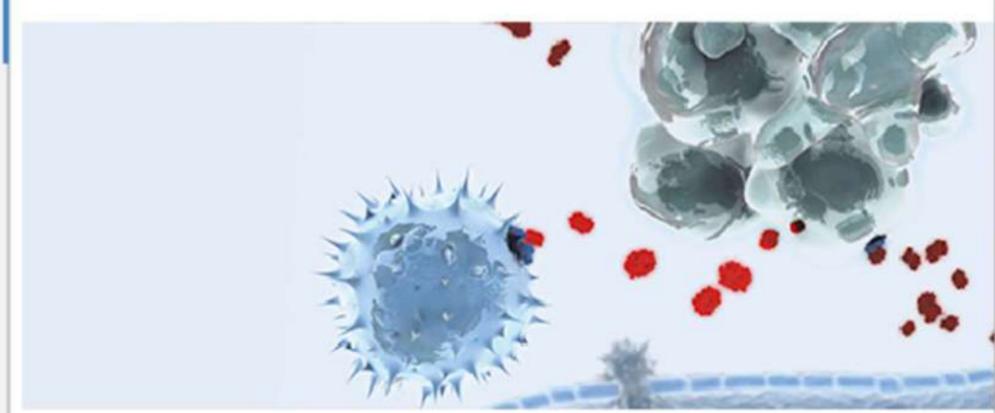
son Analyses

ter Results

A-Z Sort Refresh

- Public data resources
- Knowledge
- Genomics or proteomics data
- RNA data
- Proteomics data

- Analyzing genetic gain/loss data
- Analyzing metabolomics data
- Case studies and Support webinars
- Top help articles and FAQs
- Contacting Support
- Shortcuts



**News**

- Starting in Jan 2024, analyses >10 years old will be eligible for [automatic deletion](#).
- Sign up for an [IPA training webinar!](#)
- Check out our NEW [1-minute IPA promo video](#).
- Check out the [single cell expression tutorial](#).
- Read and watch our [gene expression tutorial with embedded videos](#).
- The IPA downtime for the upcoming **IPA 2023 Winter Release** has been scheduled for:
  - Pacific Standard Time: Friday, December 15, 5 p.m. through Sunday, December 17, 12 p.m. PST (No)
  - Central European Time: Saturday, December 16, 02:00 through Sunday, December 17, 21:00.
  - Japan Standard Time: Saturday, December 16, 10:00 through Monday, December 18, 05:00.
  - China CST (Beijing): Saturday, Saturday, December 16, 23:00 through Monday, December 18, 18:00.
- If you are new to IPA or taking a trial please see: [IPA Trial Resources](#).
- Pave your way to greatness using advanced pathway analysis: [Learn more](#).
- Read our [news](#) and sign up for our newsletter.
- Search Google Scholar for [publications that cite IPA](#).
- See how the IPA team is using Machine Learning in a [new paper](#).

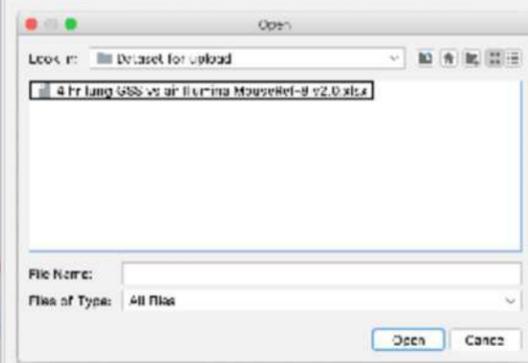
Don't Show at Startup

Dataset Upload Workflow Instructions

### Data Upload Workflow

Use Dataset Upload to import your dataset file into IPA. Once uploaded, many different analysis options exist including the Biomarker Filter, Tox and Core Analyses.

1. To upload a dataset file, [click here](#).



2. Select the dataset file (.txt, .xls, .xlsx, .csv, or .diff) from your computer and click the **Open** button.

3. Select **Flexible format** for the file format from the dropdown menu if it is not already selected.

4. Assign an **array platform** used for the dataset, if applicable. Assigning the appropriate array platform improves the accuracy of the statistics by assigning the platform as the reference set or "universe" of all possible measured molecules for that dataset. Leave as "Not specified/applicable" for RNA-seq, metabolomics, or phosphoproteomics datasets.

5. If your dataset is fairly simple (without many extraneous columns that you intend to ignore) click the **Infer Observations** button. IPA will try to automatically assign the columns for you. If this is successful, you may opt to assign additional ID columns (see step 5) or just skip to step 9. If Infer Observations is unsuccessful, just click the button again to toggle it off.

6. Assign at least one column as an **identifier column** ("ID") from the dropdown menu, then select its identifier type(s) in the secondary dropdown menu. IPA



Create New...

Search

Advanced Search

Project Manager

A-Z Sort

- My Projects
  - GWAS
    - Dataset Files
    - Analyses
      - Ulcerative Colitis-associations - 2023-10-19 11:07
      - colon cancer-association - 2023-10-18 05:52
      - Ulcerative Colitis-associations - 2023-10-18 05:19
    - Comparison Analyses
    - Biomarker Filter Results
    - Biomarker Comparison Analyses
    - MicroRNA Target Filter Results
    - BioProfiler Results
    - IsoProfiler Results
    - My Pathways
    - My Lists
  - Human Genes Chromosomal Location
  - QIAGEN Coronavirus Networks
  - Ingenuity KEGG gene lists
  - Tissue Expression
  - Example Analyses
  - Training
  - Shared Projects

Create Comparison Analysis

Select analyses for side-by-side comparison. Click View Comparison to view comparison results.

Create Comparison Analysis

Select Analyses A-Z Sort

- My Projects
  - GWAS
    - Ulcerative Colitis-associations - 2023-10-18 05:19
    - colon cancer-association - 2023-10-18 05:52
    - Ulcerative Colitis-associations - 2023-10-18 05:19
  - Shared Projects

Add >>

Analyses to Compare

- Ulcerative Colitis-associations - 2023-10-18 05:19
- colon cancer-association - 2023-10-18 05:52

Move Up ▲

Move Down ▼

<< Remove

View Comparison

Comparison Analysis Workflow Instructions

### Comparison Analyses Workflow

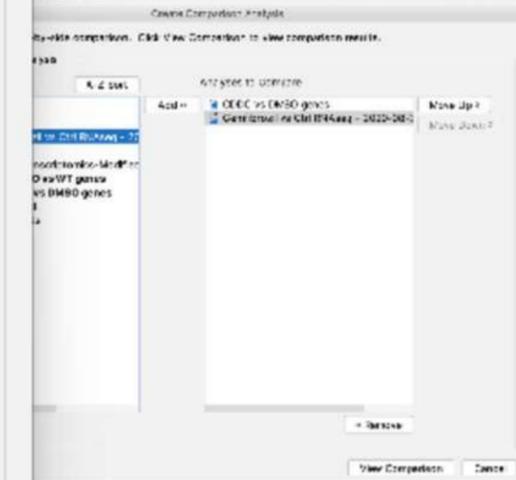
Analyses are available for Core, Tox and Metabolomics analyses.

Analysis allows you to analyze changes in biological states across multiple timepoints or dosage treatments. First run a Core, Tox or Metabolomics Analysis on your multiple projects. Then use Comparison Analysis to understand which biological processes, clinical pathology endpoints, and pathways are relevant to each timepoint or dose.

#### Run a Comparison Analysis

Comparison Analysis, [click here](#).

Projects from the left menu and click Add to add them to the Analyses to Compare window.



Comparison.

For more information, see the help manual for Comparison Analysis heatmaps for:

- [Pathways](#)
- [Regulators](#)
- [Causal Networks](#)
- [Diseases and Functions](#)

**Note:** If you do not want Workflow Instructions to open automatically: From the menu bar, select File>Preferences>Application Preferences, then deselect Show Workflow Instructions.



# Comparison Analysis – P-value



QIAGEN Land Explorer



Create New...

Genes and Chemicals Diseases and Functions Pathways and Lists Datasets and Analyses

Search

Advanced Search

Project Manager

- My Projects
  - GWAS
    - Dataset Files
    - Analyses
      - Ulcerative Colitis-association
      - colon cancer-association
      - Ulcerative Colitis-association
    - Comparison Analyses
    - Biomarker Filter Results
    - Biomarker Comparison Analyses
    - MicroRNA Target Filter Results
    - BioProfiler Results
    - IsoProfiler Results
    - My Pathways
    - My Lists
  - Human Genes Chromosomal Locations
  - QIAGEN Coronavirus Network
  - Ingenuity KEGG gene lists
  - Tissue Expression
  - Example Analyses
  - Training
  - Shared Projects

Comparison Analysis - Analysis Comparison 1

Canonical Pathways Upstream Analysis Diseases & Functions Regulator Effects Lists My Pathways Molecules Networks Comparison Settings

Settings/Legend

Filter

Measurement:  $-\log(p\text{-value})$  0.00E00  9.6

Sort Method: Score  Visualize: **p-value**

Insignificance Threshold:  (absolute value)

View Report Open Network Gene Heatmap  Expand Header

Canonical Pathw...	Ulcerative Colitis-... colon cancer-ass...
Myelination Signaling Pathway	
Human Embryonic Stem Cell Pluripotency	
Molecular Mechanisms of Cancer	
Th1 Pathway	
Role of Osteoblasts, Osteoclasts and Chondrocytes in Rheumatoid Arthritis	
BMP signaling pathway	
Th1 and Th2 Activation Pathway	
Natural Killer Cell Signaling	
Pulmonary Fibrosis Idiopathic Signaling Pathway	
Role of JAK1, JAK2 and TYK2 in Interferon Signaling	
IL-23 Signaling Pathway	
IL-12 Signaling and Production in Macrophages	
Osteoarthritis Pathway	
Cardiac Hypertrophy Signaling (Enhanced)	
IL-10 Signaling	
TGF- $\beta$ Signaling	
Cardiomyocyte Differentiation via BMP Receptors	
Interleukin-12 family signaling	

Please select a column (analysis) to display the corresponding metadata or click a heatmap ...

Save & Exit



See how the IPA team is using Machine Learning in a [new paper](#).

Don't Show at Startup

**Note:** If you do not want Workflow Instructions to open automatically. From the menu bar, select **File>Preferences>Application Preferences**, then deselect **Show Workflow Instructions**.



Create New...

Search Advanced Search

Project Manager

- My Projects
  - GWAS
    - Dataset Files
    - Analyses
      - Ulcerative Colitis-association
      - colon cancer-association
      - Ulcerative Colitis-association
    - Comparison Analyses
    - Biomarker Filter Results
    - Biomarker Comparison Analyses
    - MicroRNA Target Filter Results
    - BioProfiler Results
    - IsoProfiler Results
    - My Pathways
    - My Lists
  - Human Genes Chromosomal Locations
  - QIAGEN Coronavirus Network
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  - Tissue Expression
  - Example Analyses
  - Training
  - Shared Projects

Comparison Analysis - Analysis Comparison 1

Canonical Pathways Upstream Analysis Diseases & Functions Regulator Effects Lists My Pathways Molecules Networks Comparison Settings

Settings/Legend

Filter

Measurement:  $-\log(p\text{-value})$  0.00E00  9.6

Sort Method: **Hierarchical Clustering** Visualize: p-value

Insignificance Threshold:  (absolute value)

View Report Open Network Gene Heatmap  Expand Header  Cluster Columns

Canonical Pathway	Ulcerative Colitis-association	colon cancer-association
G alpha (12/13) signalling events		
GP6 Signaling Pathway		
Transcriptional Regulatory Network		
Senescence Pathway		
Formation of definitive endoderm		
Germ layer formation at gastrula		
Mouse Embryonic Stem Cell Pluripotency		
RHO GTPase cycle		
Signaling by TGF-beta Receptor		
Transcriptional activity of SMAD2		
RHOA Signaling		
Pancreatic Secretion Signaling Pathway		
Cell Cycle: G1/S Checkpoint Regulation		
Ephrin B Signaling		
ERK5 Signaling		
Hedgehog 'on' state		
TCF dependent signaling in response to Wnt		
MicroRNA Biogenesis Signaling Pathway		

Save & Exit

See how the IPA team is using Machine Learning in a new paper.  Don't Show at Startup

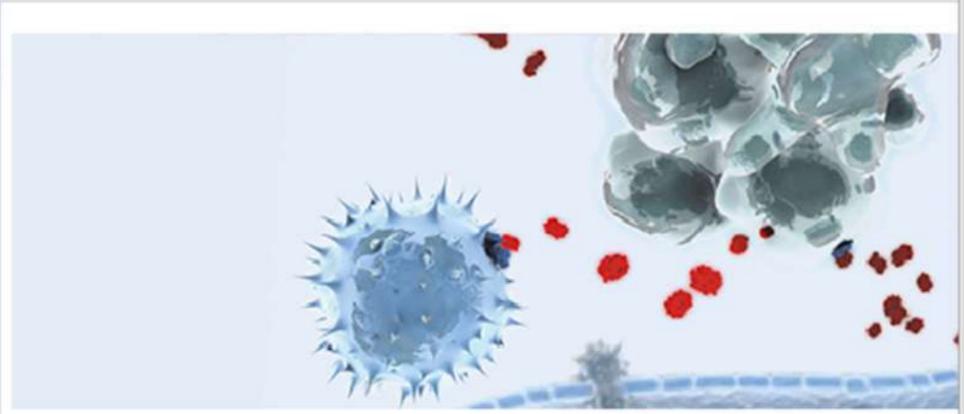
Note: If you do not want Workflow Instructions to open automatically: From the menu bar, select File>Preferences>Application Preferences, then deselect Show Workflow Instructions.



- Create New...
- Core Analysis...
- Comparison Analysis...
- Biomarker Filter...
- Biomarker Comparison Analysis...
- MicroRNA Target Filter...
- BioProfiler
- IsoProfiler
- My Pathway
- Path Designer
- Filter Dataset
- Upload Dataset...
- Advanced Search
- Project...
- Compare**
- Import Pathway

A-Z Sort Refresh

- associations - 2023-10-19 11:07
- ociation - 2023-10-18 05:52
- associations - 2023-10-18 05:19
- es
- ults
- son Analyses
- ter Results
- ublic data resources
- nowledge
- or proteomics data
- NA data
- oproteomics data



- ### News
- Starting in Jan 2024, analyses >10 years old will be eligible for [automatic deletion](#).
  - Sign up for an [IPA training webinar!](#)
  - Check out our NEW [1-minute IPA promo video](#).
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  - If you are new to IPA or taking a trial please see: [IPA Trial Resources](#).
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Don't Show at Startup

### Comparison Analysis Workflow Instructions

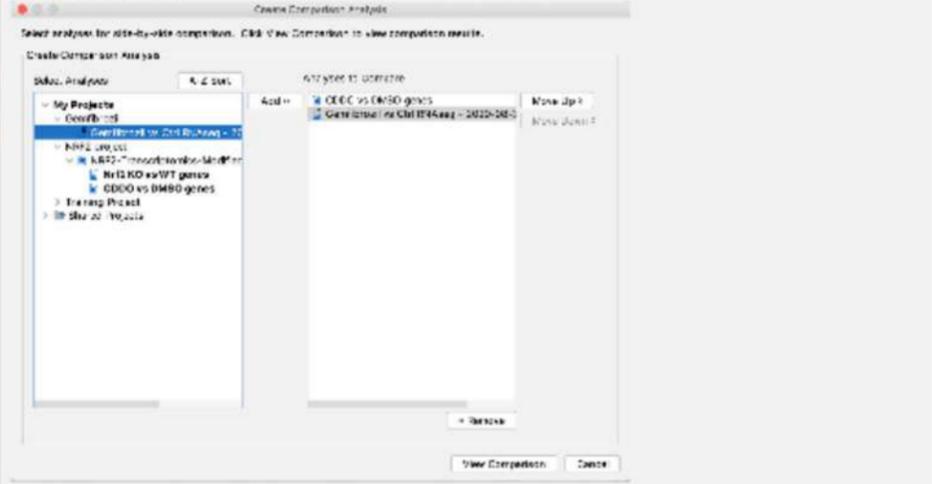
## Comparison Analyses Workflow

Comparison Analyses are available for Core, Tox and Metabolomics analyses.

Comparison Analysis allows you to analyze changes in biological states across observations. First run a Core, Tox or Metabolomics Analysis on your multiple datasets that represent timepoints or dosage treatments. Then use Comparison Analysis to understand which biological processes, clinical pathology endpoints, diseases, and pathways are relevant to each timepoint or dose.

### Create and Run a Comparison Analysis

- To start a Comparison Analysis, [click here](#).
- Choose projects from the left menu and click Add to add them to the Analyses to Compare window.



- Click View Comparison.
- To save your results, click "Save and Exit".

For more information, see the help manual for Comparison Analysis heatmaps for: [Canonical Pathways](#), [Upstream Regulators](#), [Causal Networks](#), [Diseases and Functions](#)

**Note:** If you do not want Workflow Instructions to open automatically: From the menu bar, select File>Preferences>Application Preferences, then deselect **Show Workflow Instructions**.



Create New...

Search

Advanced Search

Project Manager

### Compare

Select Entities to compare and click Add

Refresh

Calculate intersections

Add >>

<< Remove

Clear All

Click in the Venn diagram below to compare different sets. Keep the Ctrl key down to select multiple areas.

A Ulcerative Colitis-associations2 (Dataset)  
B colon cancer-association (Dataset)

Entities Comparison Results (11)

ATXN2
ERGIC1
GNA12
LOC105373831
MYRF
PITX1-AS1
PRDM1
SFMBT1
SMAD3
TET2
ZMIZ1

Add To My Pathway Add To My List Annotations

Show All

## Workflow

..., Tox and Metabolomics analyses.

changes in biological states across  
omics Analysis on your multiple  
e treatments. Then use Comparison  
cesses, clinical pathology endpoints,  
n timepoint or dose.

Click Add to add them to the Analyses to

Workflow instructions panel with a scrollable list of items and a 'New Comparison' button.

for Comparison Analysis heatmaps for:

See now the IPA team is using Machine Learning in a [new paper](#).

Don't Show at Startup

**Note:** If you do not want Workflow Instructions to open automatically: From the menu bar, select **File>Preferences>Application Preferences**, then deselect **Show Workflow Instructions**.



Project Manager Compare

Select Entities to compare and click Add

Refresh Calculate intersections

Add >> << Remove Clear All

Click in the Venn diagram below to compare different sets. Keep the Ctrl key down to select multiple areas.

A Ulcerative Colitis-associations2 (Dataset)  
B colon cancer-association (Dataset)

Entities Comparison Results (11)

ATXN2  
ERGIC1  
GNA12

### Workflow

Tox and Metabolomics analyses.

changes in biological states across omics Analysis on your multiple treatments. Then use Comparison processes, clinical pathology endpoints, timepoint or dose.

Click Add to add them to the Analyses to

Molecule Annotations

Add To My Pathway Add To My List Create Dataset Customize Table

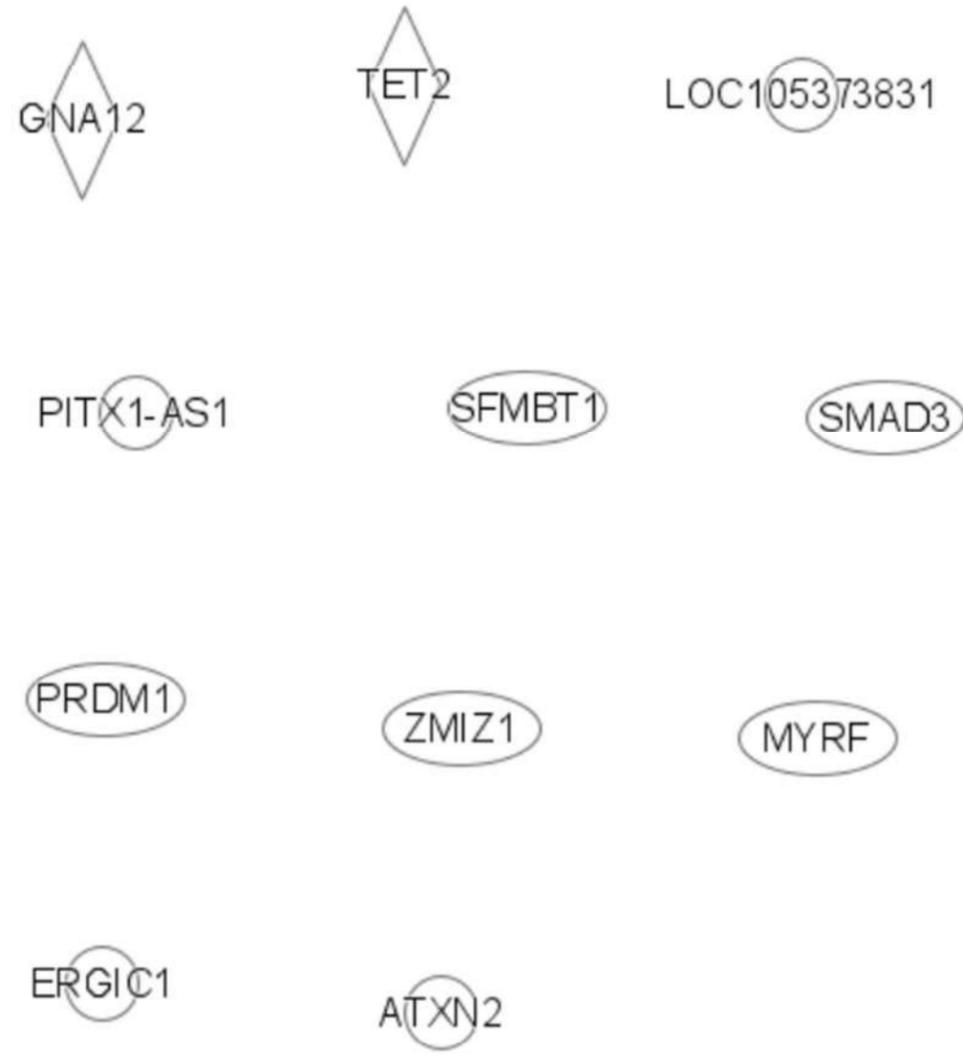
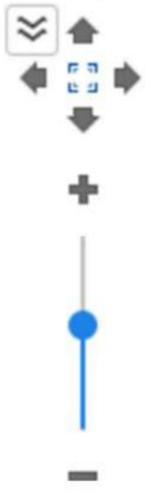
A1: Ulcerative Colitis-associations2, A2: colon cancer-association

<input checked="" type="checkbox"/>	Symbol	Entrez Gene Name	Identifier		Location	Type(s)	Biomarker Application	Drug(s)
			dbSNP/Gene Symbol	dbSNP/Gene Symbol				
<input checked="" type="checkbox"/>	ATXN2*	ataxin 2	rs653178*	rs653178	Nucleus	other		
<input checked="" type="checkbox"/>	ERGIC1*	endoplasmic reticulum-golgi...	rs564349*	rs472959	Cytoplasm	other		
<input checked="" type="checkbox"/>	GNA12*	G protein subunit alpha 12	rs1182188*	rs1182197	Plasma Membrane	enzyme	response to therapy	
<input checked="" type="checkbox"/>	LOC105373831*	uncharacterized LOC1053738...	rs17229285*	rs6434979	Other	other		
<input checked="" type="checkbox"/>	MYRF	myelin regulatory factor	rs174537	rs174537	Nucleus	transcription regu...		
<input checked="" type="checkbox"/>	PITX1-AS1*	PITX1 antisense RNA 1	rs254560*	rs647161	Other	other		
<input checked="" type="checkbox"/>	PRDM1	PR/SET domain 1	PRDM1	PRDM1	Nucleus	transcription regu...		
<input checked="" type="checkbox"/>	SFMBT1	Scm like with four mbt doma...	rs9847710	rs2581817	Nucleus	transcription regu...		

Selected/Total molecules: 11/11



New My Pathway 2



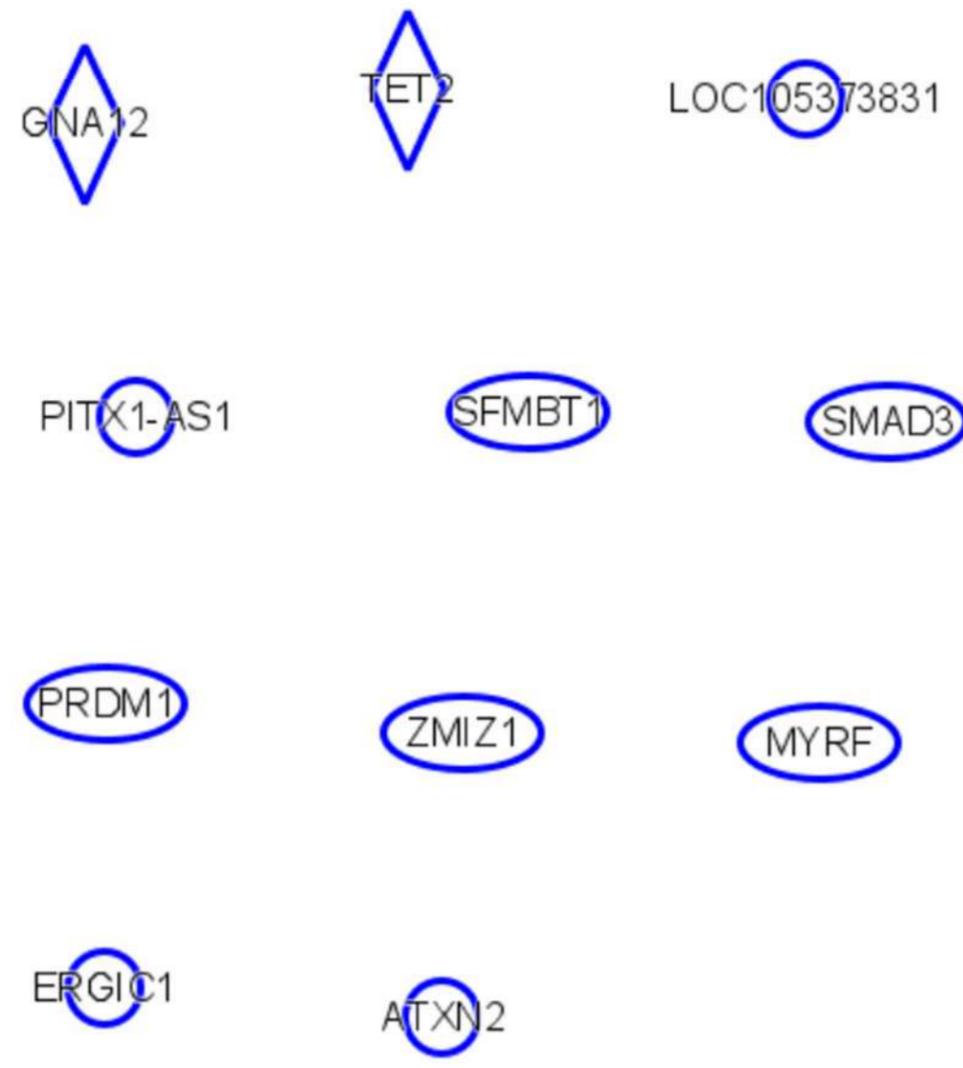
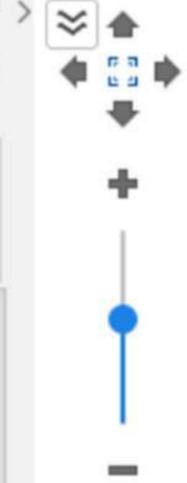
Tool: Connect

Connect selected molecules based on specified criteria. Click Apply to view new connections.

Filter Summary  
Consider all molecules and/or relationships

- General Settings
- Interactions
  - Direct
  - Indirect
- Data Sources All
- miRNA Confidence Level All
- Species All
- Tissues & Cell Lines All
- Mutation All
- Relationship Types All
- Publication Date Range All
- Node Types All
- Diseases All
- Biofluids All

New My Pathway 2



Save As Preferences Restore From Prefs Reset Apply

Tool: Connect

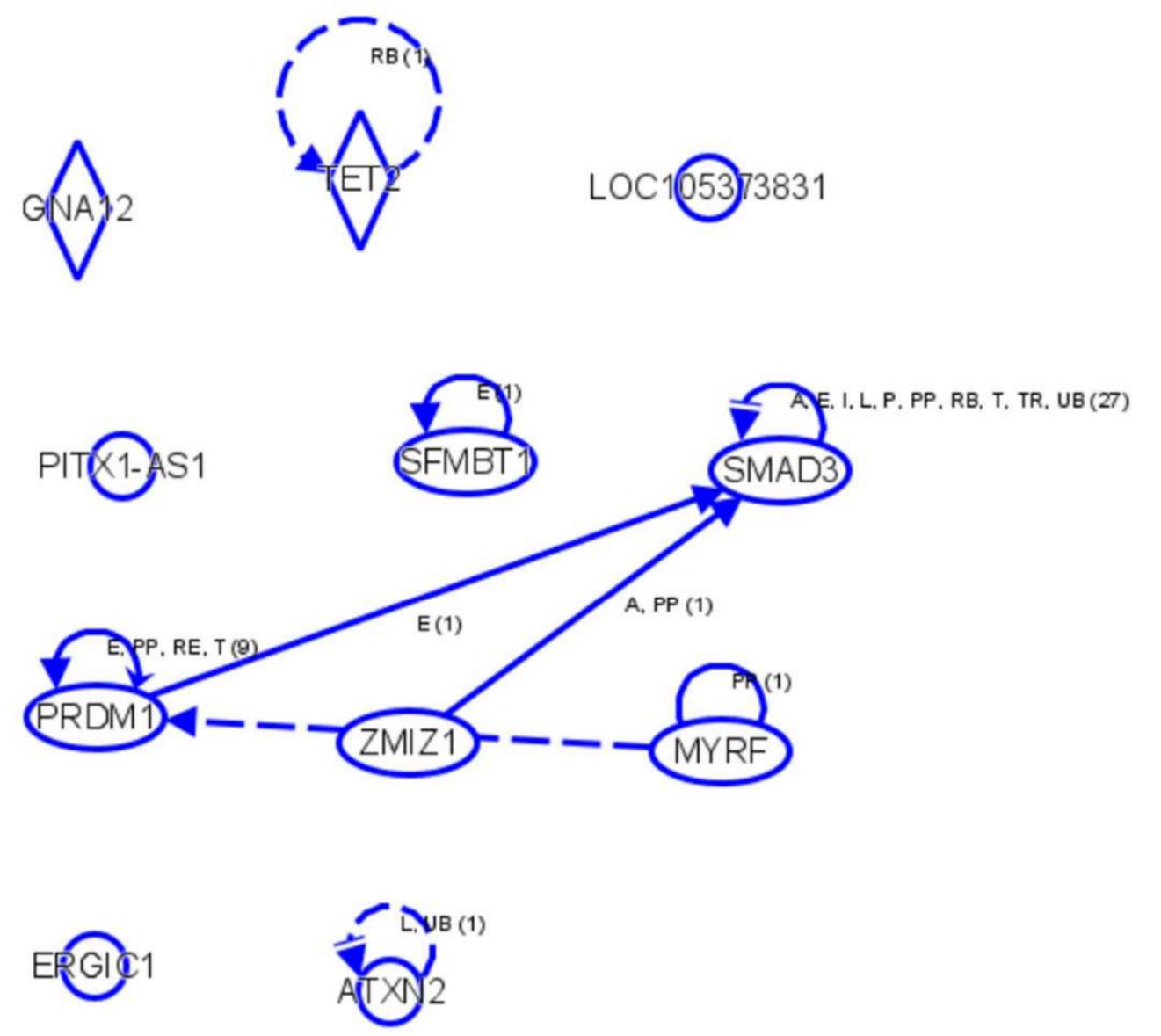
9 relationships (or 93 Findings) were added.

Filter Summary Consider all molecules and/or relationships

- General Settings
- Interactions
  - Direct
  - Indirect
- Data Sources All
- miRNA Confidence Level All
- Species All
- Tissues & Cell Lines All
- Mutation All
- Relationship Types All
- Publication Date Range All
- Node Types All
- Diseases All
- Biofluids All

Save As Preferences Restore From Prefs Reset Apply

New My Pathway 2



> [BMC Gastroenterol. 2023 Apr 27;23\(1\):136. doi: 10.1186/s12876-023-02769-5.](#)

## Peripheral blood T-lymphocyte subsets are potential biomarkers of disease severity and clinical outcomes in patients with ulcerative colitis: a retrospective study

Bailu Geng <sup>1</sup>, Xueli Ding <sup>1</sup>, Xiaoyu Li <sup>1</sup>, Hua Liu <sup>1</sup>, Wenjun Zhao <sup>1</sup>, Haihong Gong <sup>2</sup>, Zibin Tian <sup>3</sup>, Jing Guo <sup>4</sup>

Affiliations + expand

PMID: 37106335 PMCID: PMC10134527 DOI: 10.1186/s12876-023-02769-5

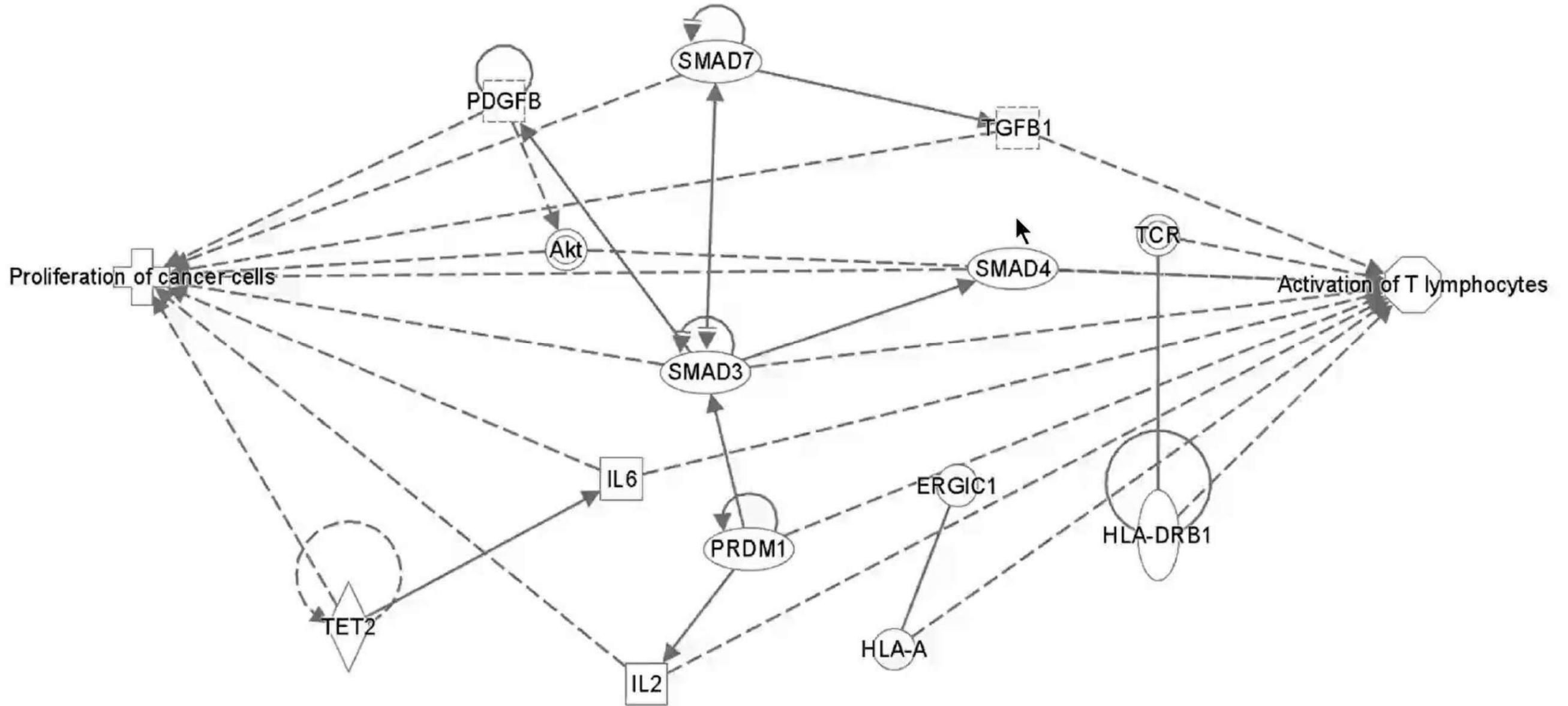
> [Am J Pathol. 2011 Apr;178\(4\):1805-13. doi: 10.1016/j.ajpath.2011.01.004.](#)

## Cytotoxic T lymphocytes efficiently recognize human colon cancer stem-like cells

Satoko Inoda <sup>1</sup>, Yoshihiko Hirohashi, Toshihiko Torigoe, Rena Morita, Akari Takahashi, Hiroko Asanuma, Munehide Nakatsugawa, Satoshi Nishizawa, Yasuaki Tamura, Tetsuhiro Tsuruma, Takeshi Terui, Toru Kondo, Kunihiko Ishitani, Tadashi Hasegawa, Koichi Hirata, Noriyuki Sato

Affiliations + expand

PMID: 21435460 PMCID: PMC3078439 DOI: 10.1016/j.ajpath.2011.01.004





Create New...

Search Advanced Search

Annotated Dataset: myList

Preview Dataset myList

Mapped IDs (16) Unmapped IDs (0) All IDs (16) Metadata

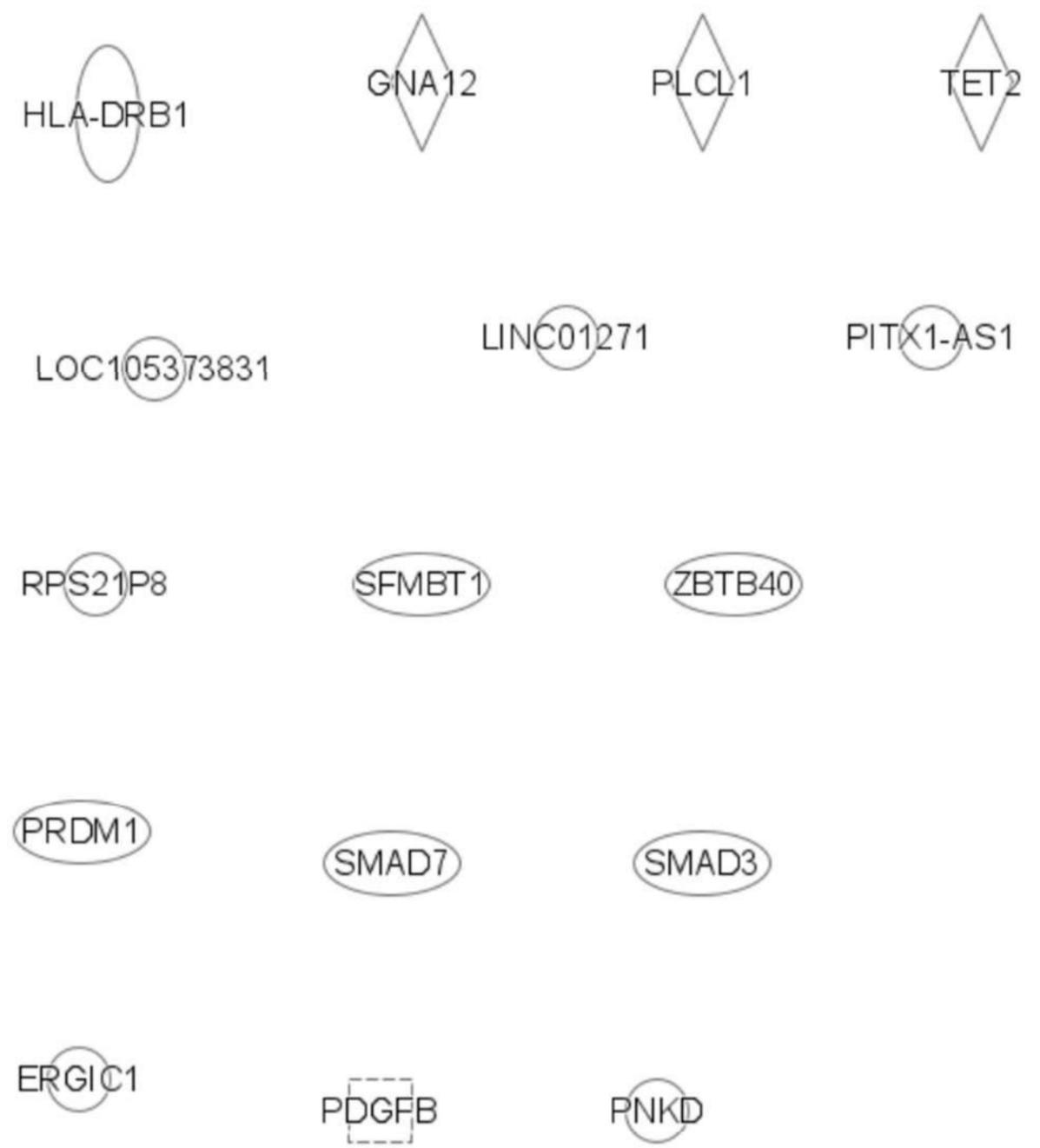
Add To My Pathway New My Pathway Create Dataset Customize Table

ID	Symbol	Entrez Gene Name	Location	Type(s)	Drug(s)
ERGIC1	ERGIC1	endoplasmic reticulum-golgi intermediate c...	Cytoplasm	other	
GNA12	GNA12	G protein subunit alpha 12	Plasma Membrane	enzyme	
HLA-DRB1	HLA-DRB1	major histocompatibility complex, class II, D...	Plasma Membrane	transmembrane receptor	apolizumab
LINC01271	LINC01271		Other	other	
LOC105373831	LOC105373831	uncharacterized LOC105373831	Other	other	
PDGFB	PDGFB	platelet derived growth factor subunit B	Extracellular Space	growth factor	dasatinib, sunitinib, pegpleranib, imatinib/...
PITX1-AS1	PITX1-AS1	PITX1 antisense RNA 1	Other	other	
PLCL1	PLCL1	phospholipase C like 1 (inactive)	Cytoplasm	enzyme	quinacrine
PNKD	PNKD	PNKD metallo-beta-lactamase domain conta...	Nucleus	other	
PRDM1	PRDM1	PR/SET domain 1	Nucleus	transcription regulator	
RPS21P8	RPS21P8	ribosomal protein S21 pseudogene 8	Other	other	
SFMBT1	SFMBT1	Scm like with four mbt domains 1	Nucleus	transcription regulator	
SMAD3	SMAD3	SMAD family member 3	Nucleus	transcription regulator	
SMAD7	SMAD7	SMAD family member 7	Nucleus	transcription regulator	
TET2	TET2	tet methylcytosine dioxygenase 2	Nucleus	enzyme	
ZBTB40	ZBTB40	zinc finger and BTB domain containing 40	Nucleus	transcription regulator	

16 / 16

Flags:  
 "D" - Duplicates. Gene/Protein/Chemical identifiers marked with an asterisk indicate that multiple identifiers in the dataset file map to a single gene/chemical in the Global Molecular Network.  
 "O" - Override molecules. Gene/Protein/Chemical identifiers marked as "Override" are displayed with italic text.  
 "A" - Gene/Protein/Chemical ID marked as Absent. The gene/protein/chemical will not be used as a focus molecule or appear in networks unless you also explicitly override this flag with the Override column.

Edit Dataset Settings Analyze/Filter Dataset Close







Create New...

activation of T lymphocytes

Search

Advanced Search

Project Manager

- My Projects
  - GWAS
    - Dataset Files
      - myList
        - Ulcerative Colitis-associations2
        - colon cancer-association
        - Ulcerative Colitis-associations
      - Analyses
        - Ulcerative Colitis-associations -
        - colon cancer-association - 2023
        - Ulcerative Colitis-associations -
      - Comparison Analyses
      - Biomarker Filter Results
      - Biomarker Comparison Analyses
      - MicroRNA Target Filter Results
      - BioProfiler Results
      - IsoProfiler Results
      - My Pathways
      - My Lists
    - Human Genes Chromosomal Location
    - QIAGEN Coronavirus Networks
    - Ingenuity KEGG gene lists

Search Results

Genes and Chemicals Diseases and Functions

Add To My Pathway New My Pathway Annotations Show Findings Effect on Function BioProfiler Activity Plot Show Functions Expand Functions

The search for activation of T lymphocytes returned 118 diseases and functions.

Matching Diseases & Functions	Associated Molecule
<input type="checkbox"/> Hematological System Development and Function	1421
<input type="checkbox"/> activation	1372
<input checked="" type="checkbox"/> Activation of T lymphocytes	1137
<input type="checkbox"/> Activation of lymphocytes	1134
<input type="checkbox"/> Activation of natural killer cells [activation of NK Lymphocyte, activation of NK cells,...]	877
<input type="checkbox"/> Activation of B lymphocytes [activation of B-enriched lymphocytes, activation of primary B lymphocytes]	262
<input type="checkbox"/> Activation of CD4+ T-lymphocytes [activation of CD4-positive T-lymphocytes, activation of CD4+ lymphocytes,...]	254
<input type="checkbox"/> Activation of CD8+ T lymphocyte [activation of CD8-positive T-lymphocytes, activation of CD8+ lymphocytes,...]	163
<input type="checkbox"/> Activation of helper T lymphocytes [activation of helper inducer T lymphocytes, activation of helper T lymphocytes,...]	130
<input type="checkbox"/> Activation of cytotoxic T cells [activation of cytotoxic T lymphocytes, activation of cytotoxic T cells,...]	88
<input type="checkbox"/> Activation of natural killer T lymphocytes	87
<input type="checkbox"/> Activation of regulatory T lymphocytes	59
<input type="checkbox"/> Activation of memory T lymphocytes	58
<input type="checkbox"/> Activation of Th1 cells [activation of Th1 lymphocytes, activation of T helper type 1 cells,...]	49
<input type="checkbox"/> Activation of naive lymphocytes	40
<input type="checkbox"/> Activation of effector lymphocytes	35
<input type="checkbox"/> Activation of naive T lymphocytes	34
<input type="checkbox"/> Activation of alpha-beta T lymphocytes [activation of alpha-beta T lymphocytes]	32
<input type="checkbox"/> Activation of mature T lymphocytes	29
<input type="checkbox"/> Activation of effector T lymphocytes	28
<input type="checkbox"/> Activation of peripheral blood lymphocytes	27
<input type="checkbox"/> Activation of Th2 cells [activation of Th2 lymphocytes, activation of th2 effector cells,...]	26
	24

Select source

Add

- Diseases or Functions
- Molecules
- Both

OK Cancel

My Pathways

New My Pathway 6

Build Overlay Path Designer Pattern Search View: Zoom: Export:

Tool: Connect

11 relationships (or 192 Findings) were added.

Filter Summary  
Consider all molecules and/or relationships

General Settings  
Interactions  
 Direct  Indirect

Data Sources All  
miRNA Confidence Level All  
Species All  
Tissues & Cell Lines All  
Mutation All  
Relationship Types All  
Publication Date Range All  
Node Types All  
Diseases All  
Biofluids All

Save As Preferences Restore From Prefs Reset Apply

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Chinese (Traditional)



Tool: Path Explorer

Add molecules to set A and B and explore shortest path(s) based on specified criteria. Click Apply to view list of shortest path(s).

Filter Summary  
Consider all molecules and/or relationships

**General Settings**

**Interactions**  
 Direct  Indirect

**Set A** Add Remove  
 ERGIC1  
 GNA12  
 HLA-DRB1

**Direction:** From Set A to Set B

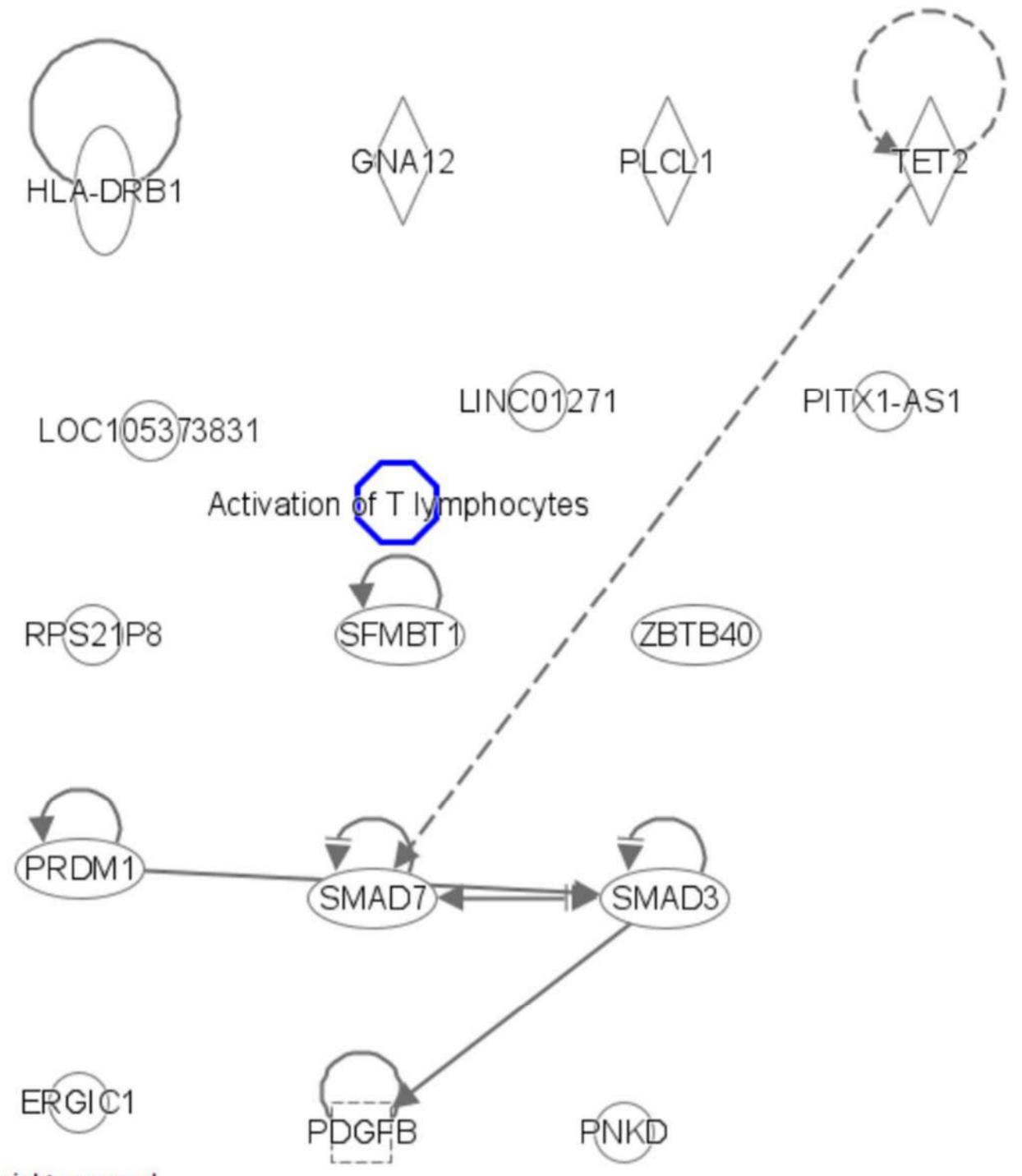
**Set B** Add Remove  
 Activation of T lymphocytes

Use Ingenuity Knowledge Base  
 Use Molecules from Analysis/Dataset/List...  
 Current Analysis/Dataset/List: None selected  
 Change Analysis/Dataset/List

Data Sources All

Save As Preferences Restore From Prefs Reset Apply

New My Pathway 6



Tool: Path Explorer

3 shortest paths were found.

Filter Summary  
 Consider all molecules and/or relationships

**General Settings**

**Interactions**  
 Direct  Indirect

**Set A** Add Remove

LINC01271  
 GNA12  
 PITX1-AS1

**Direction:** From Set A to Set B

**Set B** Add Remove

Activation of T lymphocytes

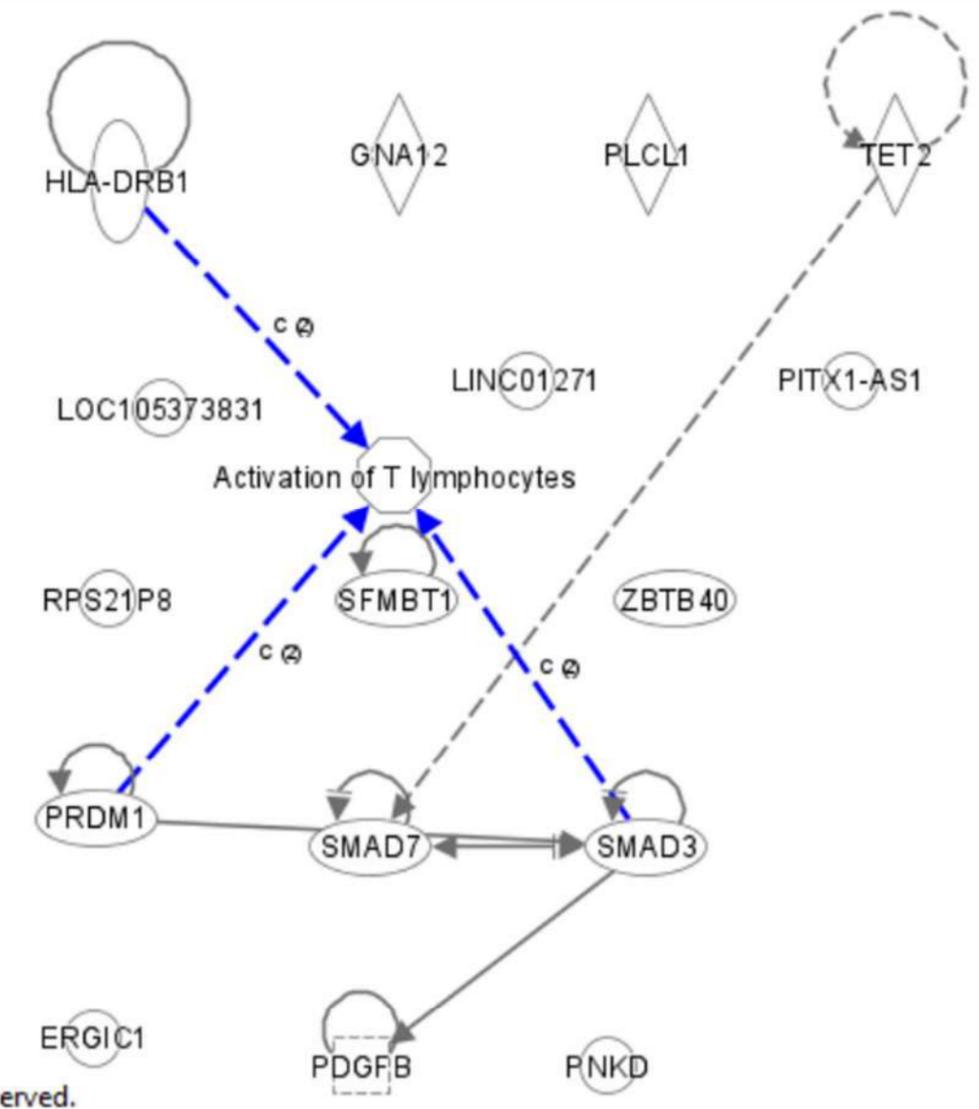
Use Ingenuity Knowledge Base  
 Use Molecules from Analysis/Dataset/List...  
 Current Analysis/Dataset/List: None selected  
 Change Analysis/Dataset/List

> Data Sources All

Save As Preferences Restore From Prefs Reset Apply

New My Pathway 6

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Add To My Pathway Highlight View Shortest Paths (3) Paths 1 - 3

Paths	Set A Molecules	Set B Molecules
<input checked="" type="checkbox"/> 1	SMAD3	Activation of T lymphocytes
<input checked="" type="checkbox"/> 2	PRDM1	Activation of T lymphocytes
<input checked="" type="checkbox"/> 3	HLA-DRB1	Activation of T lymphocytes

My Pathways

New My Pathway 6

Build Overlay Path Designer Pattern Search View: Zoom: Export:

Tool: Path Explorer

3 shortest paths were found.

Filter Summary  
Consider all molecules and/or relationships

General Settings

Interactions  
 Direct  Indirect

Set A  
 Add Remove  
 LINC01271  
 GNA12  
 PITX1-AS1

Direction: From Set A to Set B

Set B  
 Add Remove  
 Activation of T lymphocytes

Use Ingenuity Knowledge Base  
 Use Molecules from Analysis/Dataset/List...  
 Current Analysis/Dataset/List: None selected  
 Change Analysis/Dataset/List

Data Sources All

Save As Preferences Restore From Prefs Reset Apply

New My Pathway 6

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Add To My Pathway Highlight View Shortest Paths (3) Paths 1 - 3

Paths	Set A Molecules	Molecules
<input checked="" type="checkbox"/>	1	SMAD3 Activation of T lymphocytes
<input checked="" type="checkbox"/>	2	PRDM1 Activation of T lymphocytes
<input checked="" type="checkbox"/>	3	HLA-DRB1 Activation of T lymphocytes

My Pathways

☒ New My Pathway 6

Build Overlay Path Designer Pattern Search View: Zoom: Export:

Tool: Path Explorer

389 shortest paths were found.

Filter Summary  
Consider all molecules and/or relationships

**General Settings**

**Interactions**  
 Direct  Indirect

**Set A** Add Remove  
 LINC01271  
 GNA12  
 PITX1-AS1

**Direction:** From Set A to Set B --->

**Set B** Add Remove  
 Activation of T lymphocytes

Use Ingenuity Knowledge Base  
 Use Molecules from Analysis/Dataset/List...  
Current Analysis/Dataset/List: None selected  
Change Analysis/Dataset/List

> Data Sources All  
 > miRNA Confidence Level All  
 > Species All

Save As Preferences Restore From Prefs Reset Apply

**New My Pathway 6**

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Add To My Pathway Highlight View Shortest Paths + 1 (389) Paths 1 - 50

Paths	Set A Molecules	Node 1	Set B Molecules
<input checked="" type="checkbox"/> 1	SMAD3	SMAD4	Activation of T lymphocytes
<input type="checkbox"/> 2	SMAD3	IL2	Activation of T lymphocytes
<input type="checkbox"/> 3	PRDM1	IL2	Activation of T lymphocytes
<input checked="" type="checkbox"/> 4	HLA-DRB1	IL2	Activation of T lymphocytes
<input type="checkbox"/> 5	HLA-DRB1	TCR	Activation of T lymphocytes
<input checked="" type="checkbox"/> 6	PRDM1	Immunoglobulin	Activation of T lymphocytes





My Pathways

New My Pathway 6

Build Overlay Path Designer Pattern Search View: Zoom: Export:

Tool: Grow

Molecules & Canonical Pathways Diseases & Functions

Grow from selected molecules to selected diseases & functions

Indicate diseases or functions related to Any of the selected molecules

Consider all functions

Recalculate

Diseases and Functions	p-value	Molecules
Induction of actin stress fiber	3.86E-10	SMAD7, SMAD3, SM...
Proliferation of activated T lymphocytes	1.50E-09	IL2, HLA-DRB1, SMAD3...
Expansion of cancer stem cells	1.70E-09	SMAD7, SMAD3, SM...
Contractility of fibroblasts	1.70E-09	SMAD7, SMAD3, SM...
Differentiation of regulatory T lymphocytes	1.86E-09	IL2, SMAD7, HLA-DRB1...
Cytostasis of epithelial cells	2.33E-09	SMAD7, SMAD3, SM...
Activation of CD4+ T-lymphocytes	2.67E-09	IL2, HLA-DRB1, PRDM1...
Formation of abscess	3.55E-09	IL2, SMAD3, SMAD4...
Proliferation of lymphocytes	4.66E-09	IL2, SMAD7, HLA-DRB1...
Formation of skin	4.87E-09	SMAD7, HLA-DRB1, S...
Quantity of mononuclear leukocytes	5.62E-09	IL2, SMAD7, HLA-DRB1...
Differentiation of T lymphocytes	1.30E-08	IL2, SMAD7, HLA-DRB1...
Activation of T lymphocytes	1.71E-08	IL2, HLA-DRB1, SMAD3...
Transcription of DNA	3.61E-08	IL2, SMAD7, HLA-DRB1...
Differentiation of memory T lymphocytes	4.11E-08	IL2, HLA-DRB1, PRDM1...
Cell viability of lymphocytes	5.19E-08	IL2, SMAD3, PRDM1, ...
Transactivation	5.58E-08	IL2, SMAD7, SMAD3, ...
Proliferation of tumor cells	1.14E-07	IL2, SMAD7, SMAD3, ...
Quantity of T lymphocytes	1.24E-07	IL2, SMAD7, HLA-DRB1...

0/355

Reset Apply

New My Pathway 6

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Add To My Pathway Highlight

View Shortest Paths + 1 (389) Paths 1 - 50

Paths	Set A Molecules	Node 1	Set B Molecules
<input checked="" type="checkbox"/>	SMAD3	SMAD4	Activation of T lymphocytes
<input type="checkbox"/>	SMAD3	IL2	Activation of T lymphocytes
<input type="checkbox"/>	PRDM1	IL2	Activation of T lymphocytes
<input checked="" type="checkbox"/>	HLA-DRB1	IL2	Activation of T lymphocytes
<input type="checkbox"/>	HLA-DRB1	TCR	Activation of T lymphocytes
<input checked="" type="checkbox"/>	PRDM1	Immunoglobulin	Activation of T lymphocytes



Tool: Grow

Molecules & Canonical Pathways Diseases & Functions

Grow from selected molecules to selected diseases & functions

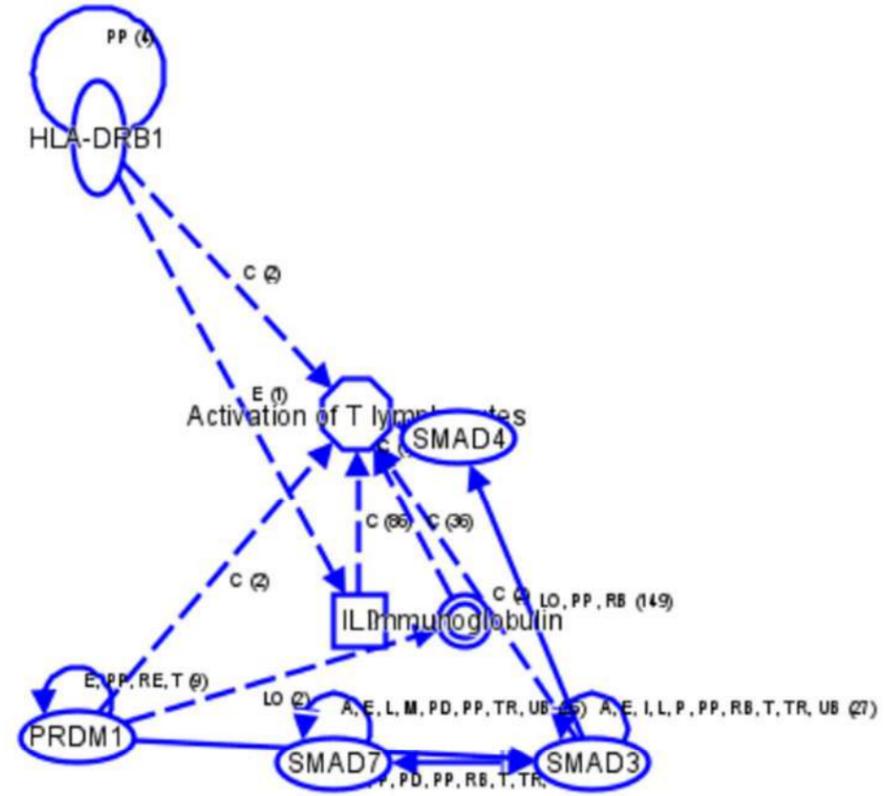
Indicate diseases or functions related to Any of the selected molecules

Consider all functions

Recalculate

Diseases and Functions	p-value	Molecules
Induction of actin stress		AD3, SM... ..all 3
Proliferation of activated		B1, SMAD3...all 5
Expansion of cancer stem		AD3, SM... ..all 3
Contractility of fibroblast		AD3, SM... ..all 3
Differentiation of regulat		HLA-DRB1...all 4
Cytostasis of epithelial ce		AD3, SM... ..all 3
Activation of CD4+ T-lympho	2.07E-05	IL2, HLA-DRB1, PRDM1...all 4
Formation of abscess	3.55E-09	IL2, SMAD3, SMAD4 ...all 3
Proliferation of lymphocytes	4.66E-09	IL2, SMAD7, HLA-DRB1...all 6
Formation of skin	4.87E-09	SMAD7, HLA-DRB1, S... ..all 5
Quantity of mononuclear leu	5.62E-09	IL2, SMAD7, HLA-DRB1 ...all 6
Differentiation of T lymphoc	1.30E-08	IL2, SMAD7, HLA-DRB1 ...all 5
Activation of T lymphocytes	1.71E-08	IL2, HLA-DRB1, SMAD3...all 5
Transcription of DNA	3.61E-08	IL2, SMAD7, HLA-DRB1...all 6
Differentiation of memory T l	4.11E-08	IL2, HLA-DRB1, PRDM1...all 3
Cell viability of lymphocytes	5.19E-08	IL2, SMAD3, PRDM1, .....all 4
Transactivation	5.58E-08	IL2, SMAD7, SMAD3, .....all 5
Proliferation of tumor cells	1.14E-07	IL2, SMAD7, SMAD3, .....all 5
Quantity of T lymphocytes	1.24E-07	IL2, SMAD7, HLA-DRB1...all 5

New My Pathway 6



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Add To My Pathway Highlight View Shortest Paths + 1 (389) Paths 1 - 50

Paths	Set A Molecules	Node 1	Set B Molecules
<input checked="" type="checkbox"/>	SMAD3	SMAD4	Activation of T lymphocytes
<input type="checkbox"/>	SMAD3	IL2	Activation of T lymphocytes
<input type="checkbox"/>	PRDM1	IL2	Activation of T lymphocytes
<input checked="" type="checkbox"/>	HLA-DRB1	IL2	Activation of T lymphocytes
<input type="checkbox"/>	HLA-DRB1	TCR	Activation of T lymphocytes
<input checked="" type="checkbox"/>	PRDM1	Immunoglobulin	Activation of T lymphocytes

Reset Apply



Tool: Grow

Molecules & Canonical Pathways Diseases & Functions

Grow from selected molecules to selected diseases & functions

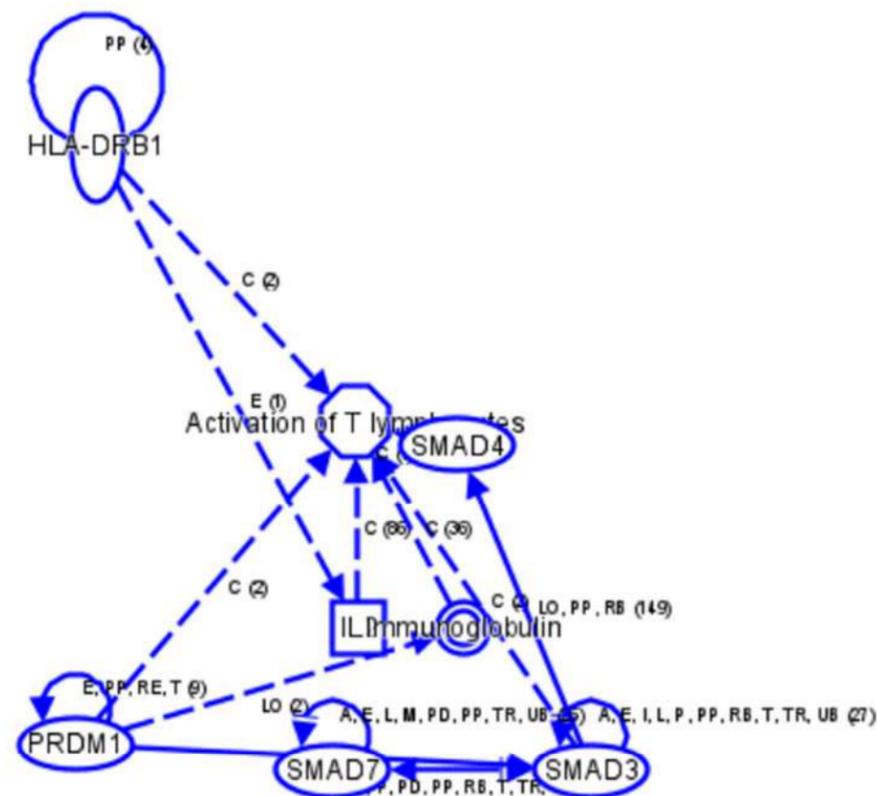
Indicate diseases or functions related to Any of the selected molecules

Consider only functions with names like proliferation\*

Recalculate

Diseases and Functions	p-value	Molecules
Proliferation of activated T lyr	1.50E-09	IL2, HLA-DRB1, SMAD3, ...all 5
Proliferation of lymphocytes	4.66E-09	IL2, SMAD7, HLA-DRB1, ...all 6
Proliferation of tumor cells	1.14E-07	IL2, SMAD7, SMAD3, ...all 5
Proliferation of beta islet cell:	3.57E-07	IL2, SMAD7, SMAD3 ...all 3
Proliferation of B lymphocyte:	4.62E-07	IL2, SMAD7, SMAD3, ...all 4
<b>Proliferation of cancer cells</b>	<b>3.63E-06</b>	<b>IL2, SMAD7, SMAD3, ...all 4</b>
Proliferation of hepatocytes	7.03E-06	SMAD7, SMAD3, SMA...all 3
Proliferation of Th1 cells	1.94E-05	IL2, HLA-DRB1 ...all 2
Proliferation of T lymphocyte:	2.47E-05	IL2, HLA-DRB1, SMAD4 ...all 3
Proliferation of hematopoieti	2.98E-05	IL2, PRDM1, SMAD4 ...all 3
Proliferation of eye cell lines	4.19E-05	SMAD3, SMAD4 ...all 2
Proliferation of splenocytes	1.25E-04	IL2, SMAD3 ...all 2
Proliferation of lymphokine a	1.47E-04	IL2 ...all 1
Proliferation of hepatic stella	1.57E-02	SMAD3 ...all 1

New My Pathway 6



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Add To My Pathway Highlight

View Shortest Paths + 1 (389) Paths 1 - 50

<input type="checkbox"/>	Paths	Set A Molecules	Node 1	Set B Molecules
<input checked="" type="checkbox"/>	1	SMAD3	SMAD4	Activation of T lymphocytes
<input type="checkbox"/>	2	SMAD3	IL2	Activation of T lymphocytes
<input type="checkbox"/>	3	PRDM1	IL2	Activation of T lymphocytes
<input checked="" type="checkbox"/>	4	HLA-DRB1	IL2	Activation of T lymphocytes
<input type="checkbox"/>	5	HLA-DRB1	TCR	Activation of T lymphocytes
<input checked="" type="checkbox"/>	6	PRDM1	Immunoglobulin	Activation of T lymphocytes

My Pathways

× New My Pathway 6

Build Overlay Path Designer Pattern Search View: Zoom: Export:

Tool: Grow

Molecules & Canonical Pathways Diseases & Functions

Grow from selected molecules to selected diseases & functions

Indicate diseases or functions related to Any of the selected molecules

Consider only functions with names like proliferation\*

Recalculate

Diseases and Functions	p-value	Molecules
Proliferation of activated T lyr	1.50E-09	IL2, HLA-DRB1, SMAD3, ...all 5
Proliferation of lymphocytes	4.66E-09	IL2, SMAD7, HLA-DRB1, ...all 6
Proliferation of tumor cells	1.14E-07	IL2, SMAD7, SMAD3, ...all 5
Proliferation of beta islet cell:	3.57E-07	IL2, SMAD7, SMAD3 ...all 3
Proliferation of B lymphocyte:	4.62E-07	IL2, SMAD7, SMAD3, ...all 4
Proliferation of cancer cells	3.63E-06	IL2, SMAD7, SMAD3, ...all 4
Proliferation of hepatocytes	7.03E-06	SMAD7, SMAD3, SMA...all 3
Proliferation of Th1 cells	1.94E-05	IL2, HLA-DRB1 ...all 2
Proliferation of T lymphocyte:	2.47E-05	IL2, HLA-DRB1, SMAD4 ...all 3
Proliferation of hematopoieti	2.98E-05	IL2, PRDM1, SMAD4 ...all 3
Proliferation of eye cell lines	4.19E-05	SMAD3, SMAD4 ...all 2
Proliferation of splenocytes	1.25E-04	IL2, SMAD3 ...all 2
Proliferation of lymphokine a	1.47E-04	IL2 ...all 1
Proliferation of hepatic stella	1.57E-02	SMAD3 ...all 1

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Add To My Pathway Highlight View Shortest Paths + 1 (389) Paths 1 - 50

Paths	Set A Molecules	Node 1	Set B Molecules
<input checked="" type="checkbox"/>	SMAD3	SMAD4	Activation of T lymphocytes
<input type="checkbox"/>	SMAD3	IL2	Activation of T lymphocytes
<input type="checkbox"/>	PRDM1	IL2	Activation of T lymphocytes
<input checked="" type="checkbox"/>	HLA-DRB1	IL2	Activation of T lymphocytes
<input type="checkbox"/>	HLA-DRB1	TCR	Activation of T lymphocytes
<input checked="" type="checkbox"/>	PRDM1	Immunoglobulin	Activation of T lymphocytes

位

My Pathways  
New My Pathway 6

Build **Overlay** Path Designer Pattern Search View: Zoom: Export:

Overlay: Biomarkers

Select Biomarker labels from table to be displayed on pathway.

Application	Disease	# Molecules	Molecule(s)
<input type="checkbox"/>	diagnosis	non-insulin-dependen...	1 IL2
<input checked="" type="checkbox"/>	diagnosis	cystic fibrosis	1 IL2
<input type="checkbox"/>	diagnosis	sarcoidosis	1 IL2
<input type="checkbox"/>	diagnosis	sarcopenia	1 IL2
<input type="checkbox"/>	diagnosis	Parkinson's disease	1 IL2
<input type="checkbox"/>	diagnosis	candidiasis	1 IL2
<input type="checkbox"/>	diagnosis	asthma	1 IL2
<input type="checkbox"/>	disease progression	Parkinson's disease	1 IL2
<input type="checkbox"/>	efficacy	HIV infection	1 IL2
<input type="checkbox"/>	efficacy	small B-cell lymphocytic ...	1 IL2
<input type="checkbox"/>	efficacy	leukemia	1 IL2
<input type="checkbox"/>	efficacy	lymphoma	1 IL2
<input type="checkbox"/>	efficacy	multiple myeloma	1 IL2
<input type="checkbox"/>	efficacy	obesity	1 IL2
<input type="checkbox"/>	efficacy	non-Hodgkin lymphoma	1 IL2
<input type="checkbox"/>	efficacy	rheumatoid arthritis	1 IL2
<input type="checkbox"/>	efficacy	myeloproliferative synd...	1 IL2
<input type="checkbox"/>	efficacy	acute myeloid leukemia	1 IL2
<input type="checkbox"/>	efficacy	myelodysplastic syndro...	1 IL2
<input type="checkbox"/>	efficacy	marginal zone cell lym...	1 IL2
<input type="checkbox"/>	efficacy	follicular lymphoma	1 IL2
<input type="checkbox"/>	efficacy	breast cancer	1 IL2
<input type="checkbox"/>	prognosis	colorectal cancer	1 SMAD4
<input type="checkbox"/>	response to therapy	rheumatoid arthritis	1 IL2
<input type="checkbox"/>	unspecified applic...	rheumatoid arthritis	1 SMAD4
<input type="checkbox"/>	unspecified applic...	gastric mucosa-associa...	1 Immunoglobulin
<input type="checkbox"/>	unspecified applic...	ovarian cancer	1 Immunoglobulin
<input type="checkbox"/>	unspecified applic...	colorectal carcinoma	1 SMAD4

HLA-DRB1

Activation of T lymphocytes

IL2

Immunoglobulin

PRDM1

SMAD7

SMAD3

SMAD4

Proliferation of cancer cells

BM diagnosis - cystic fibrosis

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Add To My Pathway Highlight View Shortest Paths + 1 (389) Paths 1 - 50

Paths	Set A Molecules	Node 1	Set B Molecules
<input checked="" type="checkbox"/>	1 SMAD3	SMAD4	Activation of T lymphocytes
<input type="checkbox"/>	2 SMAD3	IL2	Activation of T lymphocytes
<input type="checkbox"/>	3 PRDM1	IL2	Activation of T lymphocytes
<input checked="" type="checkbox"/>	4 HLA-DRB1	IL2	Activation of T lymphocytes
<input type="checkbox"/>	5 HLA-DRB1	TCR	Activation of T lymphocytes
<input checked="" type="checkbox"/>	6 PRDM1	Immunoglobulin	Activation of T lymphocytes

Mode Label Interactive OFF

My Pathways

New My Pathway 6

Build **Overlay** Path Designer Pattern Search View: [View Icon]

Overlay: **Analyses, Datasets & Lists**

Clear  Ignore Analysis Cutoff

Overlay datasets, analyses and lists

Index Name (select to overlay)

Please add a dataset, analysis or list.

**Add...**

Matching molecules

Symbol Display name

Show node charts for multi overlay:  Always  For rows selected above

Graph overlay options

Display isoform badges on nodes

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Add To My Pathway Highlight

Paths Set A Molecules Set B Molecules

<input checked="" type="checkbox"/>	1	SMAD3	SMAD4	Activation of T lymphocytes
<input type="checkbox"/>	2	SMAD3	IL2	Activation of T lymphocytes
<input type="checkbox"/>	3	PRDM1	IL2	Activation of T lymphocytes
<input checked="" type="checkbox"/>	4	HLA-DRB1	IL2	Activation of T lymphocytes
<input type="checkbox"/>	5	HLA-DRB1	TCR	Activation of T lymphocytes
<input checked="" type="checkbox"/>	6	PRDM1	Immunoglobulin	Activation of T lymphocytes

HLA-DRB1

Activation of T lymphocytes

SMAD4

IL2

PRDM1

SMAD7

IL2

BM diagnosis - cyst

Select a dataset, analysis or list

Please manually select analyses, datasets, and lists to overlay, or use search below

A-Z Sort  Search Refresh

- My Projects
  - GWAS
  - Human Genes Chromosomal Location
  - QIAGEN Coronavirus Networks
  - Ingenuity KEGG gene lists
  - Tissue Expression
  - Example Analyses
  - Training
  - Shared Projects

Use search (above) for Analysis Match datasets and analyses

OK Cancel



ulcerative colitis

Search

Examples: liver, "mouse OR rat", "HeLa NOT 3T3", "p?3", "ovar\*"

Search Results

Showing first 1779 results out of 1779 in 29233ms for query [ulcerative colitis]

Folder Types

- [dataset \(973\)](#)
- [analysis \(806\)](#)

Projects

- [HumanDisease \(1348\)](#)

Open Add to Overlay Customize Table

Creation Date: 2023/... - 2023/... (1/45)

Name	Matching Term	Type	Creation Date
1- ulcerative colitis (UC) [colonic mucosa] 26910	exp_meta_data.control.dis...	analysis	2023/09/30 23:22
1- ulcerative colitis (UC) [colonic mucosa] 33570	exp_meta_data.case.disea...	analysis	2023/09/30 23:22
1- ulcerative colitis (UC) [colon] 10226	exp_meta_data.control.dis...	analysis	2023/09/30 23:22
1- ulcerative colitis (UC) [colonic mucosa] 16417	exp_meta_data.compariso...	analysis	2023/09/30 23:21
1- ulcerative colitis (UC) [ileal mucosa] 10425	exp_meta_data.case.disea...	analysis	2023/09/30 23:21
1- ulcerative colitis (UC) [intestinal organoid] 1527	exp_meta_data.compariso...	analysis	2023/09/30 23:21
1- ulcerative colitis (UC) [colonic mucosa] 17871	exp_meta_data.compariso...	analysis	2023/09/30 23:20
1- ulcerative colitis (UC) [mesenteric adipose tissue] 25716	exp_meta_data.compariso...	analysis	2023/09/30 23:20
1- ulcerative colitis (UC) [colonic mucosa] 19885	exp_meta_data.compariso...	analysis	2023/09/30 23:20
1- ulcerative colitis (UC) [pouch mucosa] 26938	exp_meta_data.compariso...	analysis	2023/09/30 23:20
1- ulcerative colitis (UC) [colonic mucosa] 25110	exp_meta_data.compariso...	analysis	2023/09/30 23:19
1- ulcerative colitis (UC) [rectum] 8043	exp_meta_data.compariso...	analysis	2023/09/30 23:19
1- ulcerative colitis (UC) [colonic mucosa] 26770	exp_meta_data.compariso...	analysis	2023/09/30 23:19
1- ulcerative colitis (UC) [sigmoid colon] 19333	exp_meta_data.control.dis...	analysis	2023/09/30 23:19
1- ulcerative colitis (UC) [colonic mucosa] 29017	exp_meta_data.compariso...	analysis	2023/09/30 23:18
1- ulcerative colitis (UC) [peripheral blood] 14819	exp_meta_data.compariso...	analysis	2023/09/30 23:18
1- ulcerative colitis (UC) [peripheral blood] 154	exp_meta_data.control.dis...	analysis	2023/09/30 23:17
1- ulcerative colitis (UC) [rectal mucosa] 2363	exp_meta_data.compariso...	analysis	2023/09/30 23:17

Libraries > OmicSoft > DiseaseLand > HumanDisease > Analyses

1- ulcerative colitis (UC) [colon] 10226

Case/Control Differences

Key	Case	Control
response	sensitive	resistant

Comparison Context

comparisoncategory **Responder vs. Non-Responder**

comparisoncontrast **TNFIadequateResponder:Response => NA -> sensitive vs resistant**

diseasestate **ulcerative colitis (UC)**

dosage **100 mg**

organism **human**

platformname **NGS.Illumina.HiSeq2000**

subjecttreatment **etrolizumab**

tissue **colon**

treatmentstatus **none**

All Experiment Metadata

case.diseasestate **ulcerative colitis (UC)**

case.dosage **100 mg**

case.response **sensitive**

case.sampleids **GSM1872906;GSM1872912;GSM1872913;GSM187...**



ulcerative colitis

Search

Examples: liver, "mouse OR rat", "HeLa NOT 3T3", "p?3", "ovar\*"

Search Results

Showing first 1779 results out of 1779 in 29233ms for query [ulcerative colitis]

Folder Types

- [dataset \(973\)](#)
- [analysis \(806\)](#)

Projects

- [HumanDisease \(1348\)](#)

Open

Add to Overlay

Customize Table

Creation Date: 2023/... - 2023/... (1/45)

Name	Matching Term	Type	Creation Date
1- ulcerative colitis (UC) [colonic mucosa] 26910	exp_meta_data.control.dis...	analysis	2023/09/30 23:22
1- ulcerative colitis (UC) [colonic mucosa] 33570	exp_meta_data.case.disea...	analysis	2023/09/30 23:22
1- ulcerative colitis (UC) [colon] 10226	exp_meta_data.control.dis...	analysis	2023/09/30 23:22
1- ulcerative colitis (UC) [colonic mucosa] 16417	exp_meta_data.compariso...	analysis	2023/09/30 23:21
1- ulcerative colitis (UC) [ileal mucosa] 10425	exp_meta_data.case.disea...	analysis	2023/09/30 23:21
1- ulcerative colitis (UC) [intestinal organoid] 1527	exp_meta_data.compariso...	analysis	2023/09/30 23:21
1- ulcerative colitis (UC) [colonic mucosa] 17871	exp_meta_data.compariso...	analysis	2023/09/30 23:20
1- ulcerative colitis (UC) [mesenteric adipose tissue] 25716	exp_meta_data.compariso...	analysis	2023/09/30 23:20
1- ulcerative colitis (UC) [colonic mucosa] 19885	exp_meta_data.compariso...	analysis	2023/09/30 23:20
1- ulcerative colitis (UC) [pouch mucosa] 26938	exp_meta_data.compariso...	analysis	2023/09/30 23:20
1- ulcerative colitis (UC) [colonic mucosa] 25110	exp_meta_data.compariso...	analysis	2023/09/30 23:19

Analyses to overlay

1- ulcerative colitis (UC) [colon] 10226

Up ^

Down v

Remove

Clear All

Overlay Now

Libraries > OmicSoft > DiseaseLand > HumanDisease > Analyses

1- ulcerative colitis (UC) [colon] 10226

Case/Control Differences

Key	Case	Control
response	sensitive	resistant

Comparison Context

comparisoncategory **Responder vs. Non-Responder**

comparisoncontrast **TNFIadequateResponder:Response => NA -> sensitive vs resistant**

diseasestate **ulcerative colitis (UC)**

dosage **100 mg**

organism **human**

platformname **NGS.Illumina.HiSeq2000**

subjecttreatment **etrolizumab**

tissue **colon**

treatmentstatus **none**

All Experiment Metadata

case.diseasestate **ulcerative colitis (UC)**

case.dosage **100 mg**

case.response **sensitive**

case.sampleids **GSM1872906;GSM1872912;GSM1872913;GSM187...**

My Pathways

New My Pathway 6

Build Overlay Path Designer Pattern Search View: Zoom: Export:

Overlay: Analyses, Datasets & Lists

Clear  Ignore Analysis Cutoff

Overlay datasets, analyses and lists

Index	Name (select to overlay)	Matching
1	1- ulcerative colitis (UC) [colon] 10226	8

Add more...

Matching molecules

Symbol	Display name	Index	Measurement	Expr Log Ratio
IGHV1-3	Immunoglobulin parent node	1	1- ulcerative colitis (UC)	2.374 ↑
IGHV2-26	Immunoglobulin parent node	1	1- ulcerative colitis (UC)	1.560 ↑
IGHV3-48	Immunoglobulin	1	1- ulcerative colitis (UC)	1.536 ↑

Show node charts for multi overlay:  Always  For rows selected above

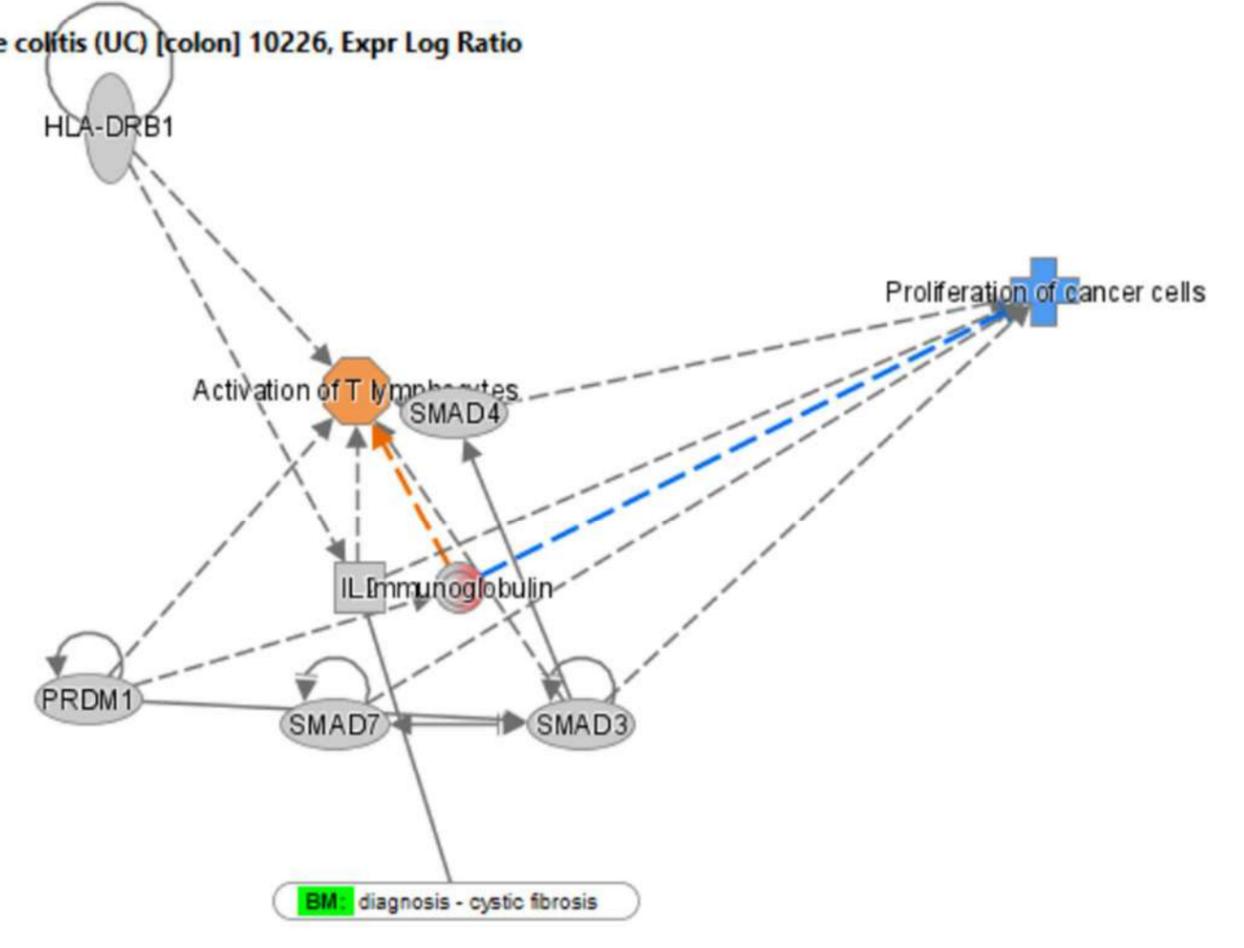
Graph overlay options

Fill	Measurement	Show value	Range
<input checked="" type="radio"/> Expr Log Ratio	<input type="checkbox"/>	<input type="checkbox"/>	-1.641  2.088
<input type="radio"/> Expr p-value	<input type="checkbox"/>	<input type="checkbox"/>	
<input checked="" type="checkbox"/> Display isoform badges on nodes			

New My Pathway 6

Overlay: 1- ulcerative colitis (UC) [colon] 10226, Expr Log Ratio

Show Legend



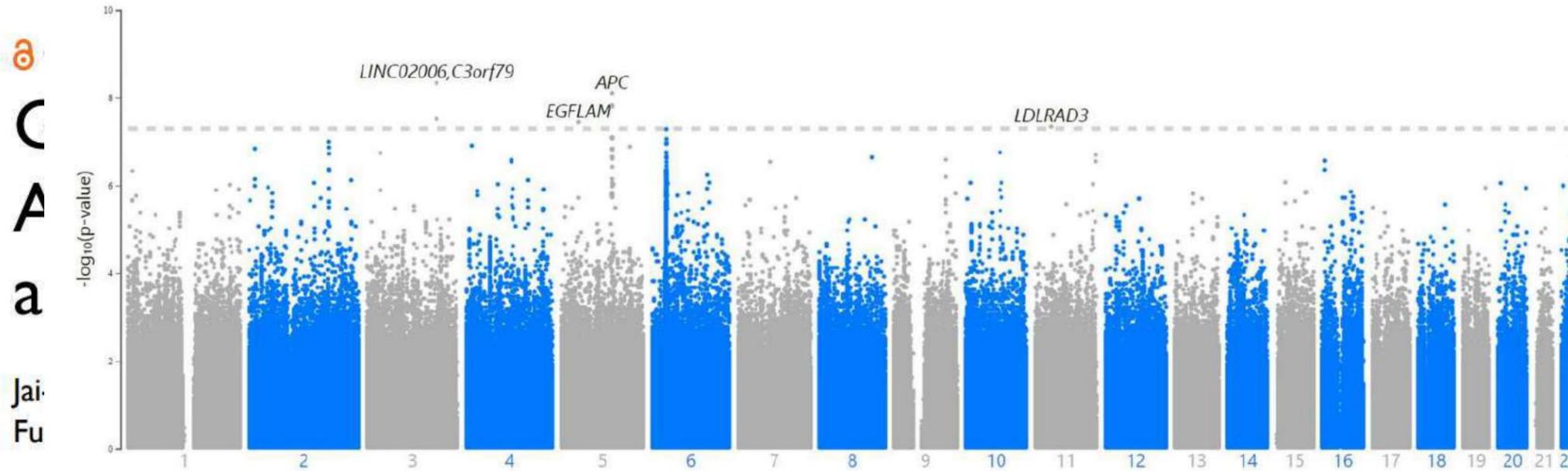
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Add To My Pathway Highlight View Shortest Paths + 1 (389) Paths 1 - 50

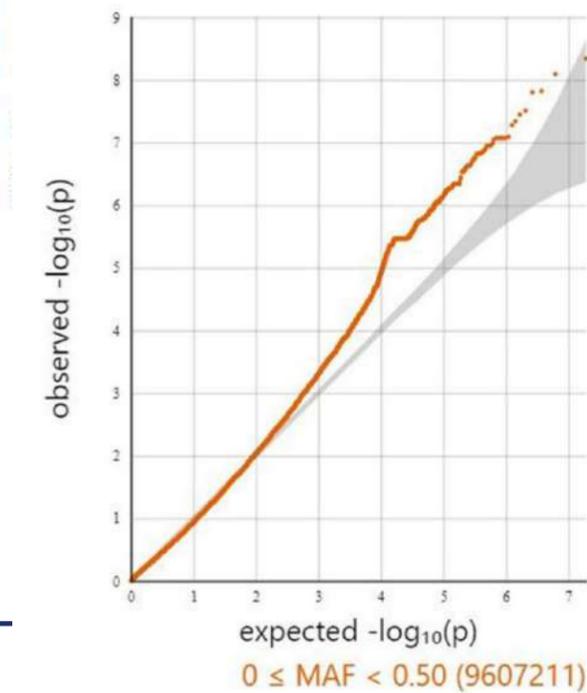
Paths	Set A Molecules	Node 1	Set B Molecules
<input checked="" type="checkbox"/> 1	SMAD3	SMAD4	Activation of T lymphocytes
<input type="checkbox"/> 2	SMAD3	IL2	Activation of T lymphocytes
<input type="checkbox"/> 3	PRDM1	IL2	Activation of T lymphocytes
<input checked="" type="checkbox"/> 4	HLA-DRB1	IL2	Activation of T lymphocytes
<input type="checkbox"/> 5	HLA-DRB1	TCR	Activation of T lymphocytes
<input checked="" type="checkbox"/> 6	PRDM1	Immunoglobulin	Activation of T lymphocytes

# Clinical, Cosmetic and Investigational Dermatology

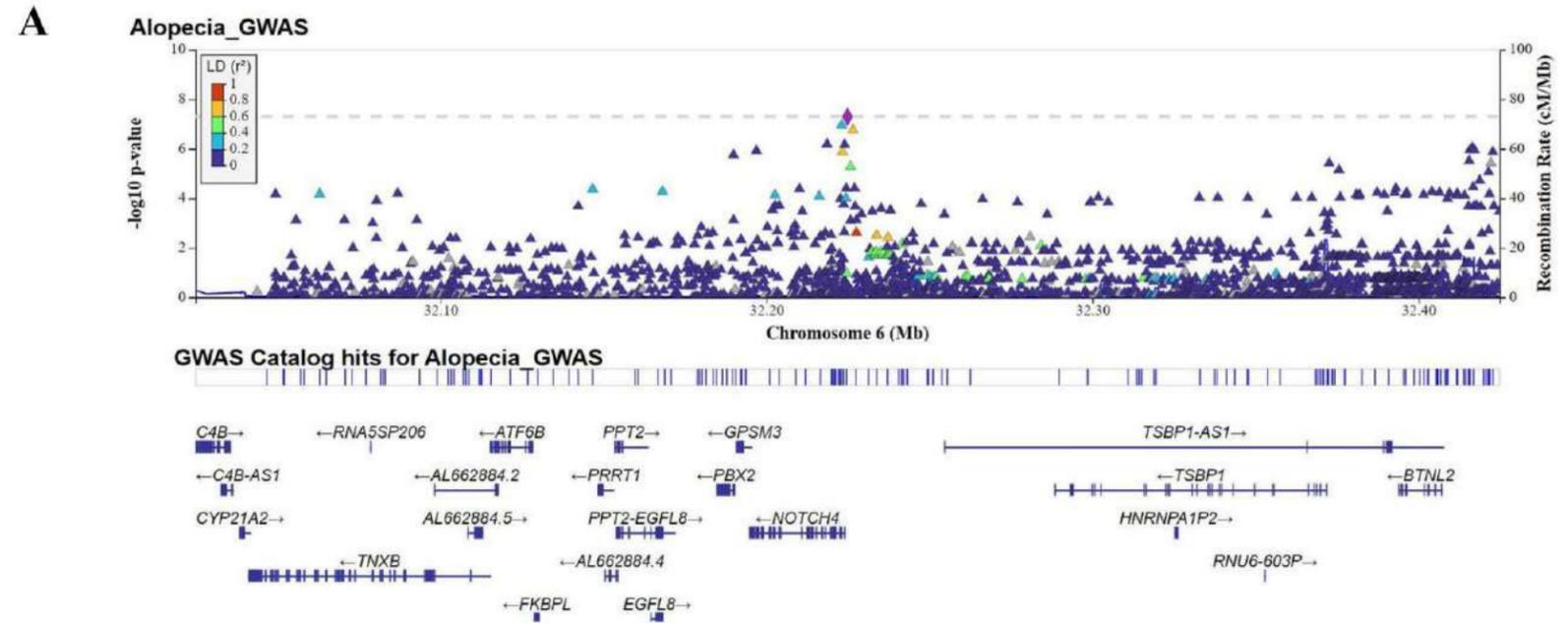
# Dovepress



<sup>1</sup>Department of Medical Research, China Medical University Hospital, China Medical University, Taichung, 404327, Taiwan; <sup>2</sup>Million-Person Precision Medicine Initiative, Department of Medical Research, China Medical University Hospital, Taichung, 404327, Taiwan; <sup>3</sup>Department of Biological Science and Technology, China Medical University, Taichung, 406040, Taiwan; <sup>4</sup>Division of Plastic and Reconstructive Surgery, Department of Surgery, Taipei

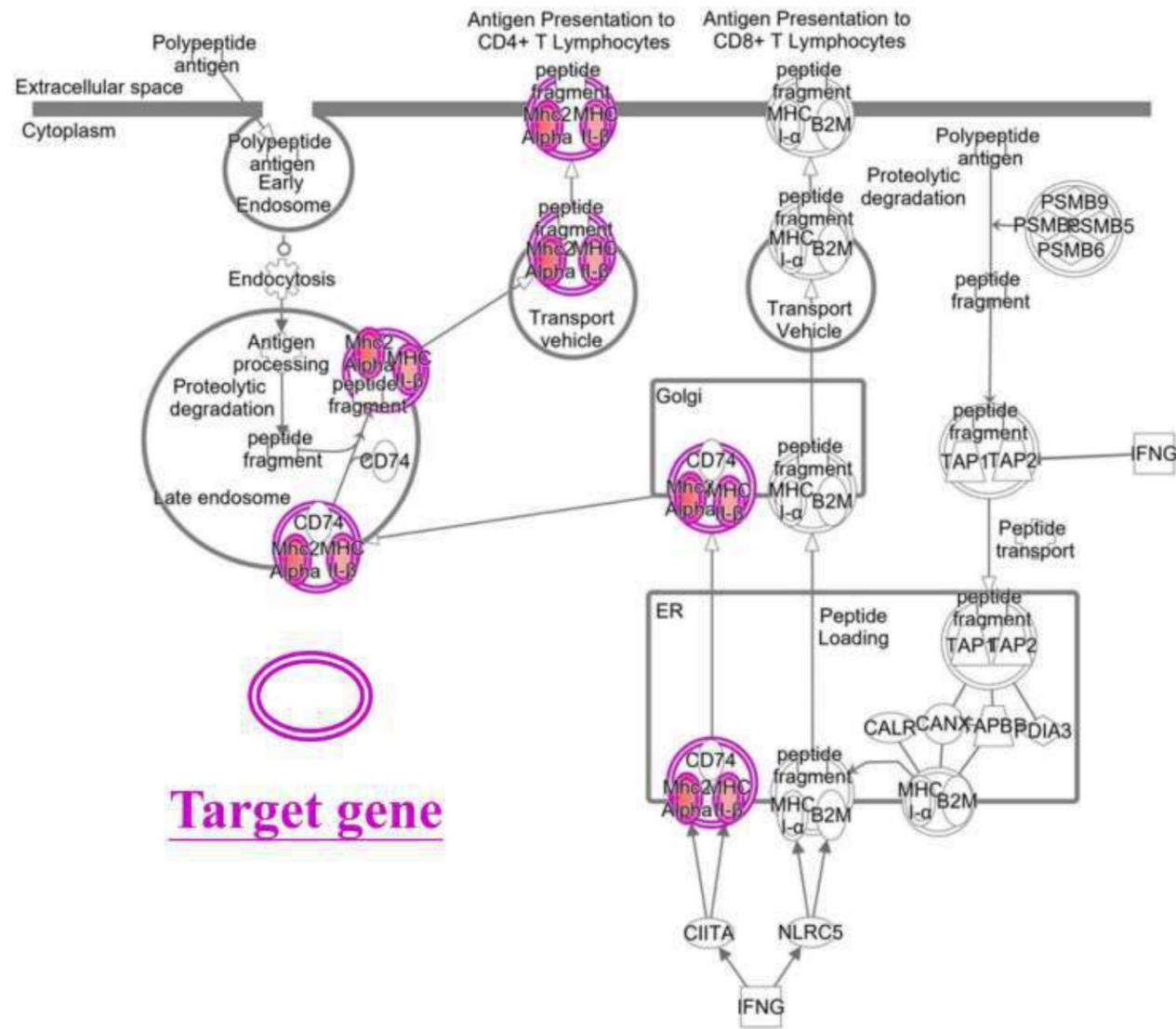


<sup>5</sup>Department of C...  
<sup>6</sup>...  
<sup>7</sup>...  
<sup>8</sup>...



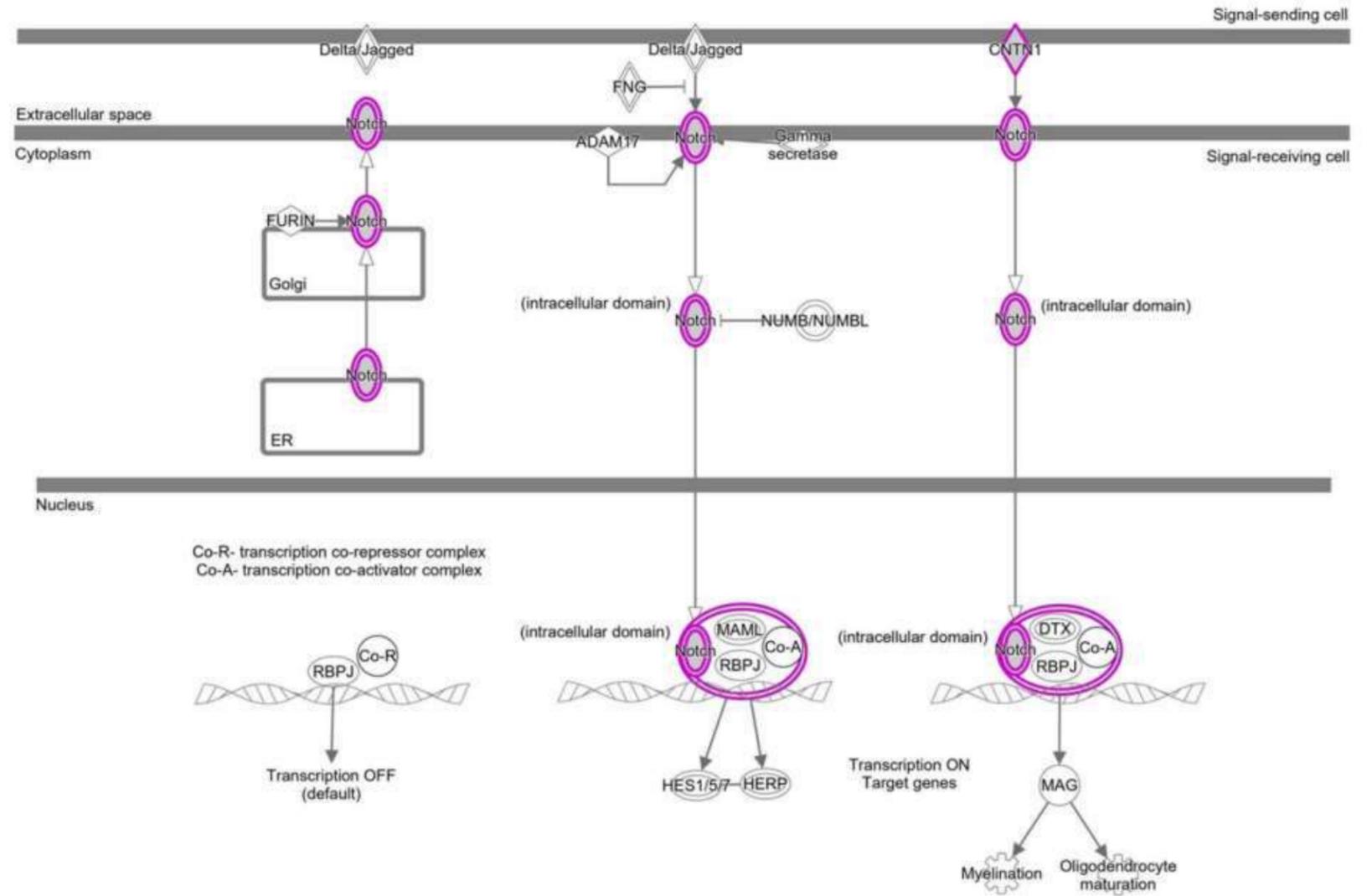


**A**



### Antigen Presentation Pathway

**B**



### Notch Signaling

**Figure 5** Network analyses of the antigen presentation pathway target gene (**A**) and the gene that Notch signaling targets (**B**). IPA software analyzed network analysis (SNPs gene loci,  $P < 1E-05$ ).



## Better Care with Better Knowledge

若有需要進一步的資訊或在使用軟體上遇到問題歡迎聯繫以下窗口：  
周儀柔 資深業務專員 [GraceChou@gga.asia](mailto:GraceChou@gga.asia) 02-2795 1777 #3019  
林文賢 經理 [BryceLin@gga.asia](mailto:BryceLin@gga.asia) 02-2795 1777 #3665

[Bioinfo@GGA.ASIA](mailto:Bioinfo@GGA.ASIA)