



Introduction to **Cytoscape**

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www.cranfield.ac.uk



Why networks?

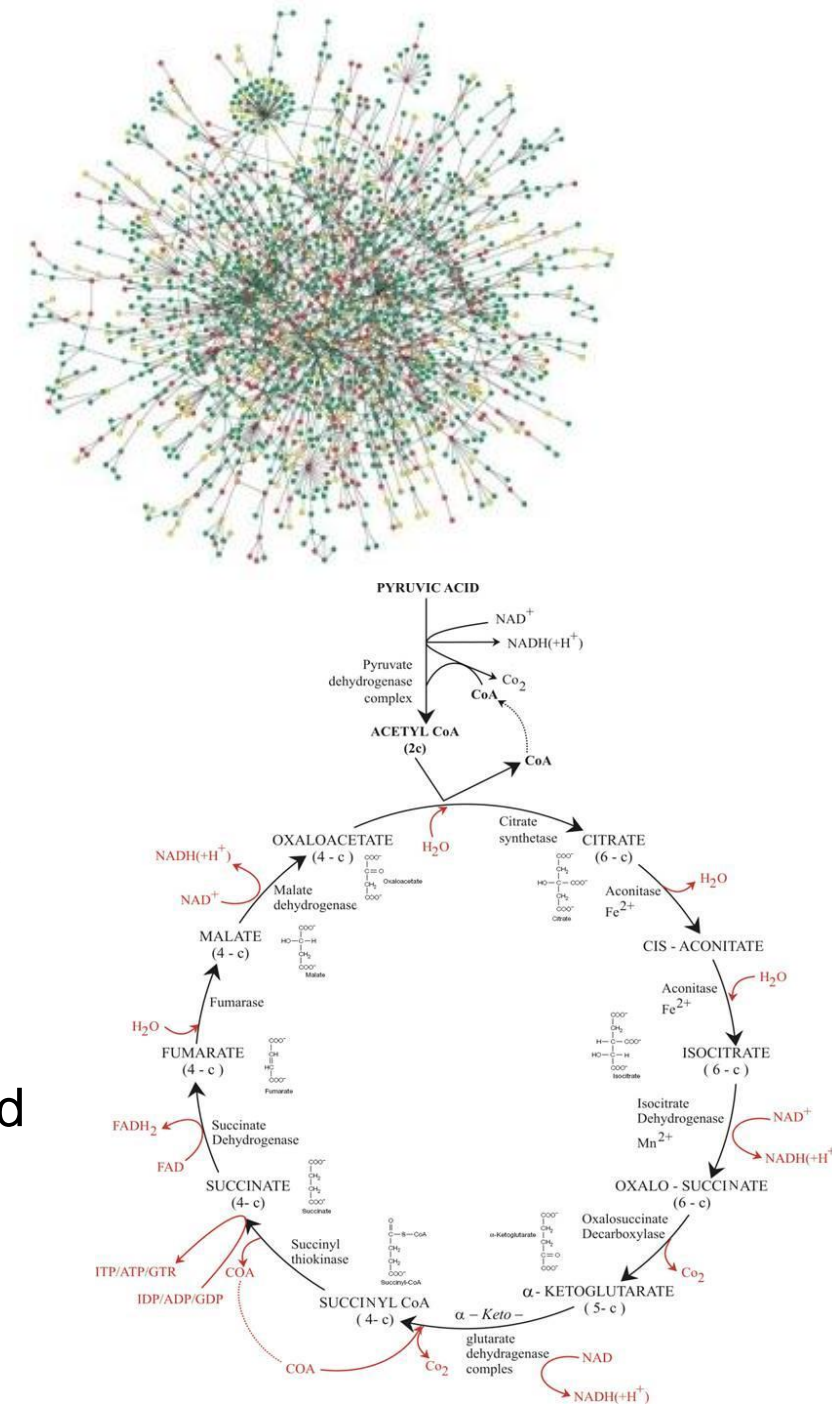
- Complex systems are better described as networks of interacting components
- The topology of a network characterises the underlying complex system (global topology parameters) and its individual components (local topology parameters)
- Network topology parameters are easily compared
- Useful for discovering patterns in large data sets
- Allow the integration of multiple data types

Two main types of networks in biology:

- Interaction networks (protein-protein - PPI, protein-ligand, protein-transcript...) - molecular interactions between proteins, or between proteins and chemicals e.g. MINT, Intact
- Pathways (regulatory, signaling, metabolic...) - Sequence of enzymatic or other reactions by which one biological material is converted to another e.g. Biocarta, Reactome, Biopax

Also:

- Association networks – genes and proteins often associated in the literature or in expression studies
- Author networks – who works with whom!





Networks as graphs

Formal **graph definition**: A graph G is a pair of two sets V (nodes) and E (edges): $G = (V, E)$

Neighbors are two nodes n_1 and n_2 connected by an edge

Neighborhood is the set of all neighbors of node n

Connectivity k_n is the size of the neighborhood of n

Degree k is the number of edges incident on n

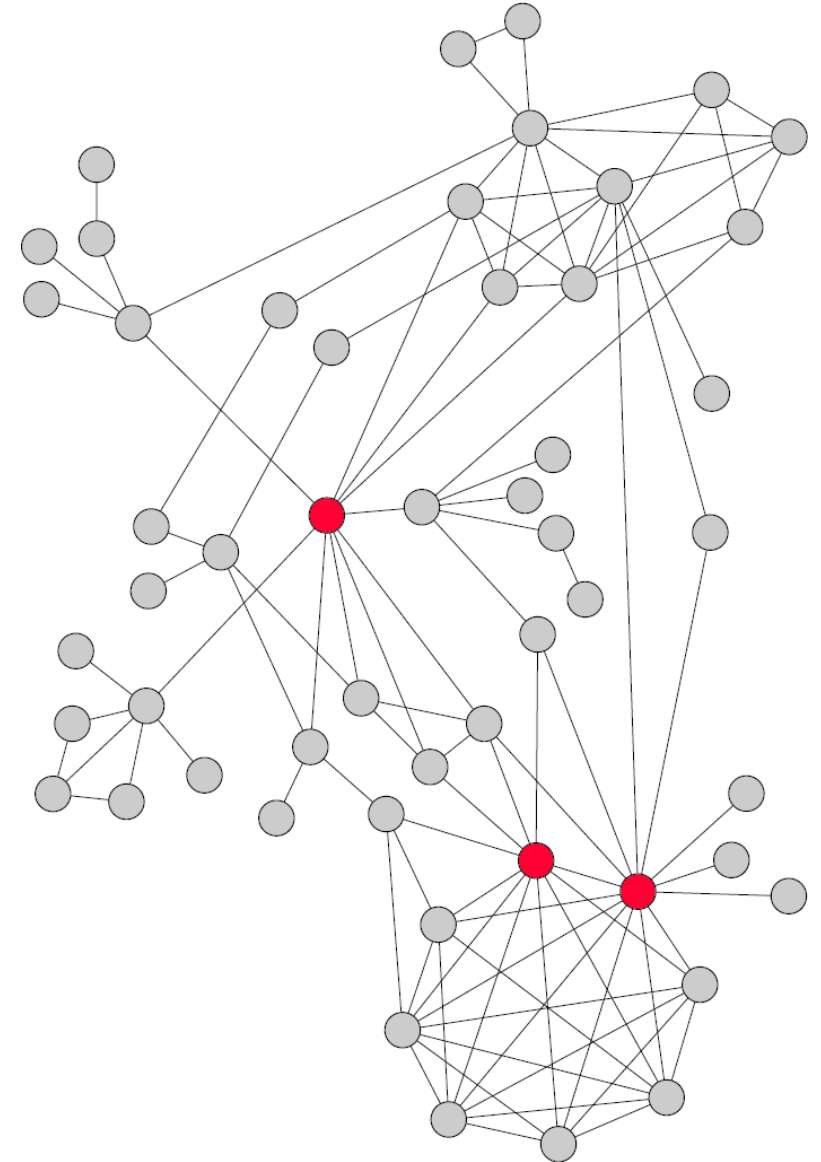
Node degree and shortest path

A **hub** is a node with an exceptionally high degree, larger than the average node degree (see red nodes).

A **shortest path** between the nodes n and m is a path between n and m of minimal length.

The shortest path length, or **distance**, between n and m is the length of a shortest path between n and m .

The **characteristic path length** is the average shortest path length, the expected distance between two connected nodes.



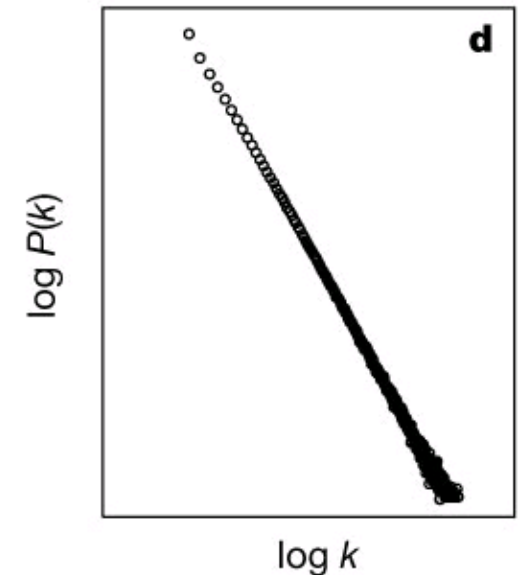
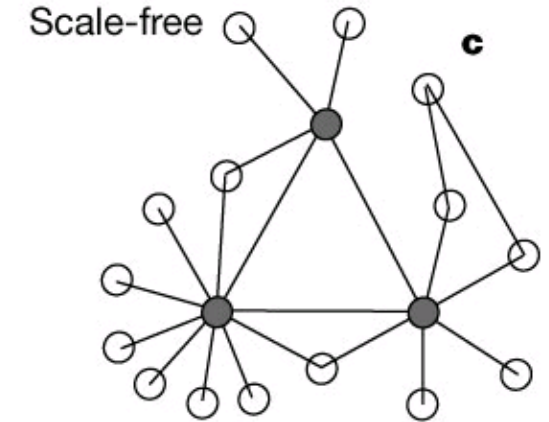
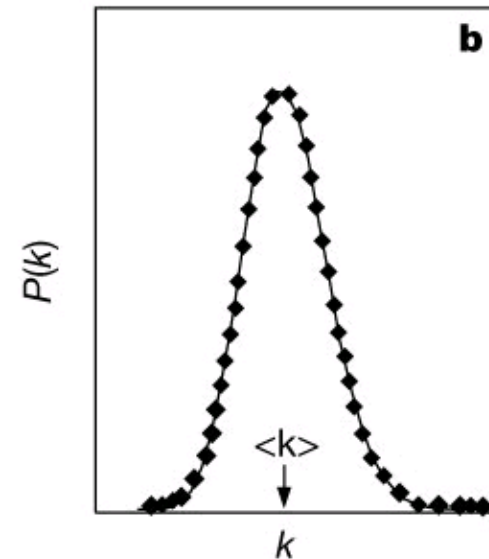
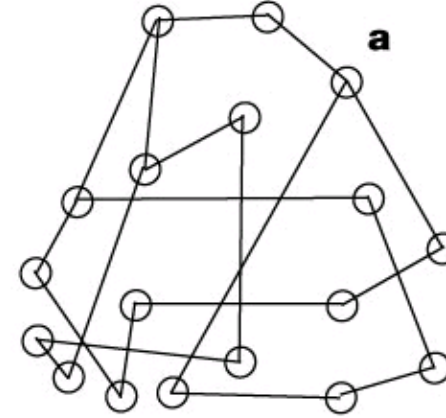
Random networks vs. scale-free networks

Random networks are homogeneous, most nodes have the same number of links:

- not robust to arbitrary node failure

Scale-free networks have a number of highly connected nodes:

- robust to random failure
- very sensitive to hub failures





Cytoscape

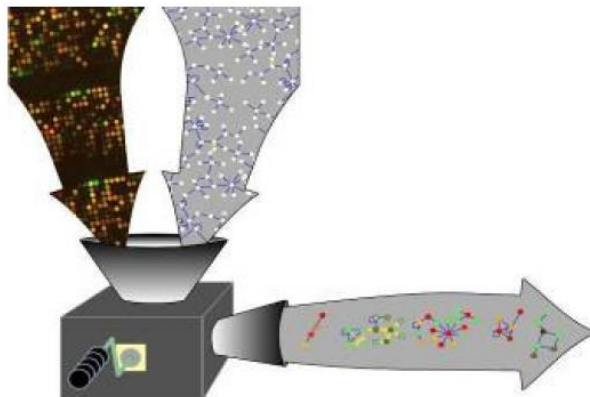
Open source software for visualising, integrating, and analysing biological data.

Main interface:

Data importing and visualisation

Advanced parsing allowing the association of data from many different formats

Complex layout customisation and automation options



Ideker and Lauffenburger, *Trends in Biotech.* (2003) -
Cited from UCSD presentation 18th January

More advanced: Apps

Clustering

Enrichment analysis

Gene expression analysis

Gene function prediction

Network evolution

...

Consortium



Institute for Systems Biology



University of California at San Diego



Memorial Sloan-Kettering Cancer Center



Institut Pasteur



Agilent Technologies



University of California at San Francisco



University of Toronto



University of Michigan



Unilever



Data tables

Nodes and edges can have additional data associated with them and stored in tables.

- Annotation
- Gene ontology
- Mass spectrometry data
- Gene expression data
- Identifiers from different databases

Filters allow you to search for nodes or edges based on that data.

It is also available to analysis apps.

The screenshot shows a software interface for network analysis. The top menu bar includes File, Edit, View, Select, Layout, Apps, Tools, and Help. Below the menu is a toolbar with various icons for file operations, network manipulation, and search. The main window is titled "Session: New Session". On the left, a "Control Panel" shows a list of networks, with "RUAL.subset.sif" selected. The central area displays a network graph with blue nodes and edges. On the right, a "Table Panel" shows a table of data for the selected network. The table has three columns: "shared name", "name", and "Official HUGO Symbol". The row for "TP53" (ID 7157) is highlighted. At the bottom, there are tabs for "Node Table", "Edge Table", and "Network Table". A "Memory" indicator is visible in the bottom right corner.

shared name	name	Official HUGO Symbol
9013	9013	TAF1C
6285	6285	S100B
3428	3428	IFI16
2908	2908	NR3C1
598	598	BCL2L1
3313	3313	HSPA9B
7157	7157	TP53
4792	4792	NFKBIA
2068	2068	ERCC2
9112	9112	MTA1
467	467	ATF3
204851	204851	HIPK1
2071	2071	ERCC3
10474	10474	TADA3L
3148	3148	HMGB2
4601	4601	NCL



Data import

Import Columns From Table

Target Table Data

Where to Import Table Data: To a Network Collection

Select a Network Collection

Network Collection: RUAL.subset...

Import Data as: Node Table Columns

Key Column for Network: shared na...

Case Sensitive Key Values: ☒

Preview

Click on a column to edit it.

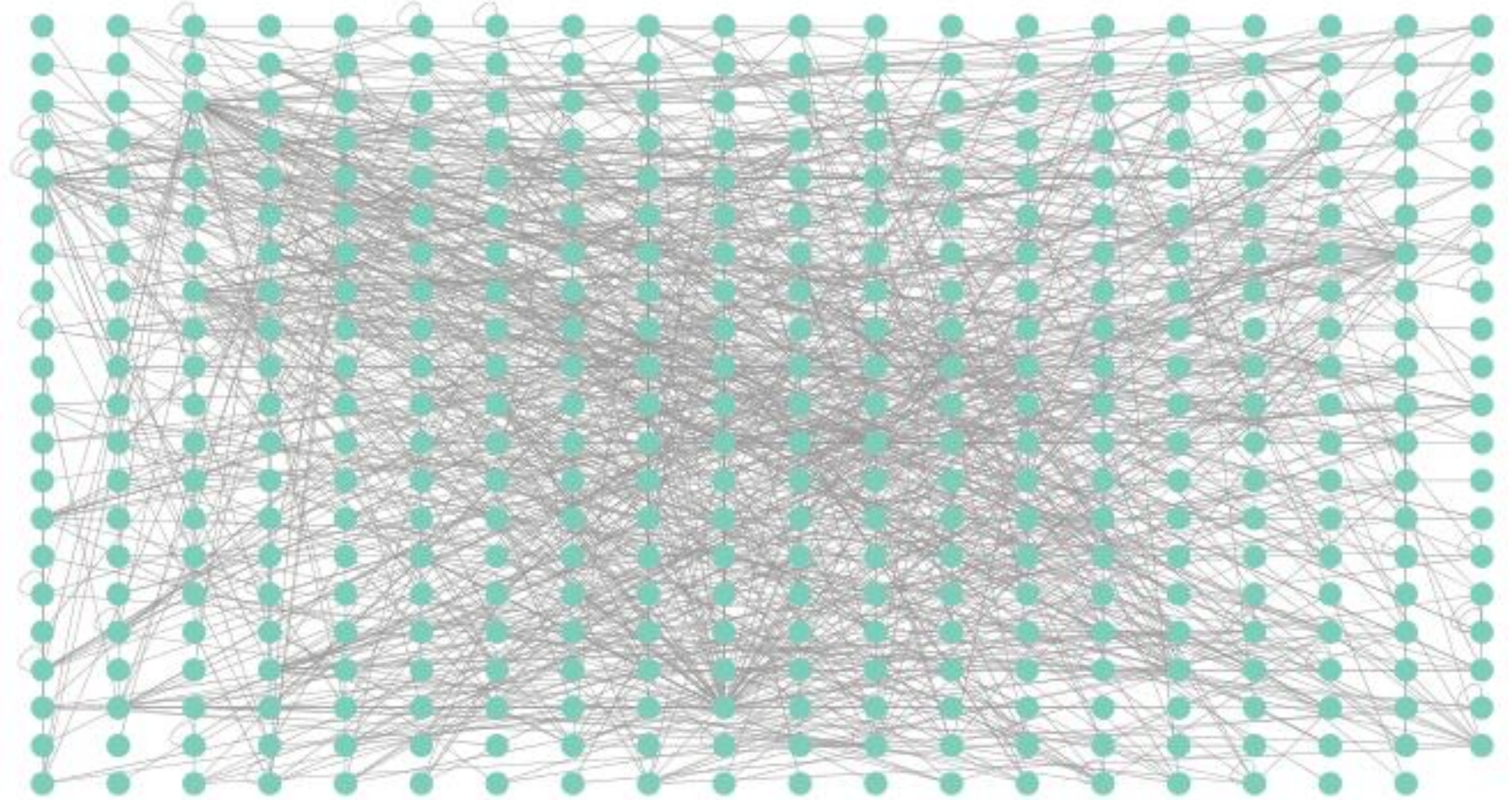
Select All Select None

Id	=	Official HUGO Symbol
26003	=	GORASP2
81628	=	TSC22D4
9729	=	KIAA0408
55722	=	Cep72
56658	=	TRIM39
56658	=	TRIM39
7186	=	TRAF2
83461	=	CDCA3

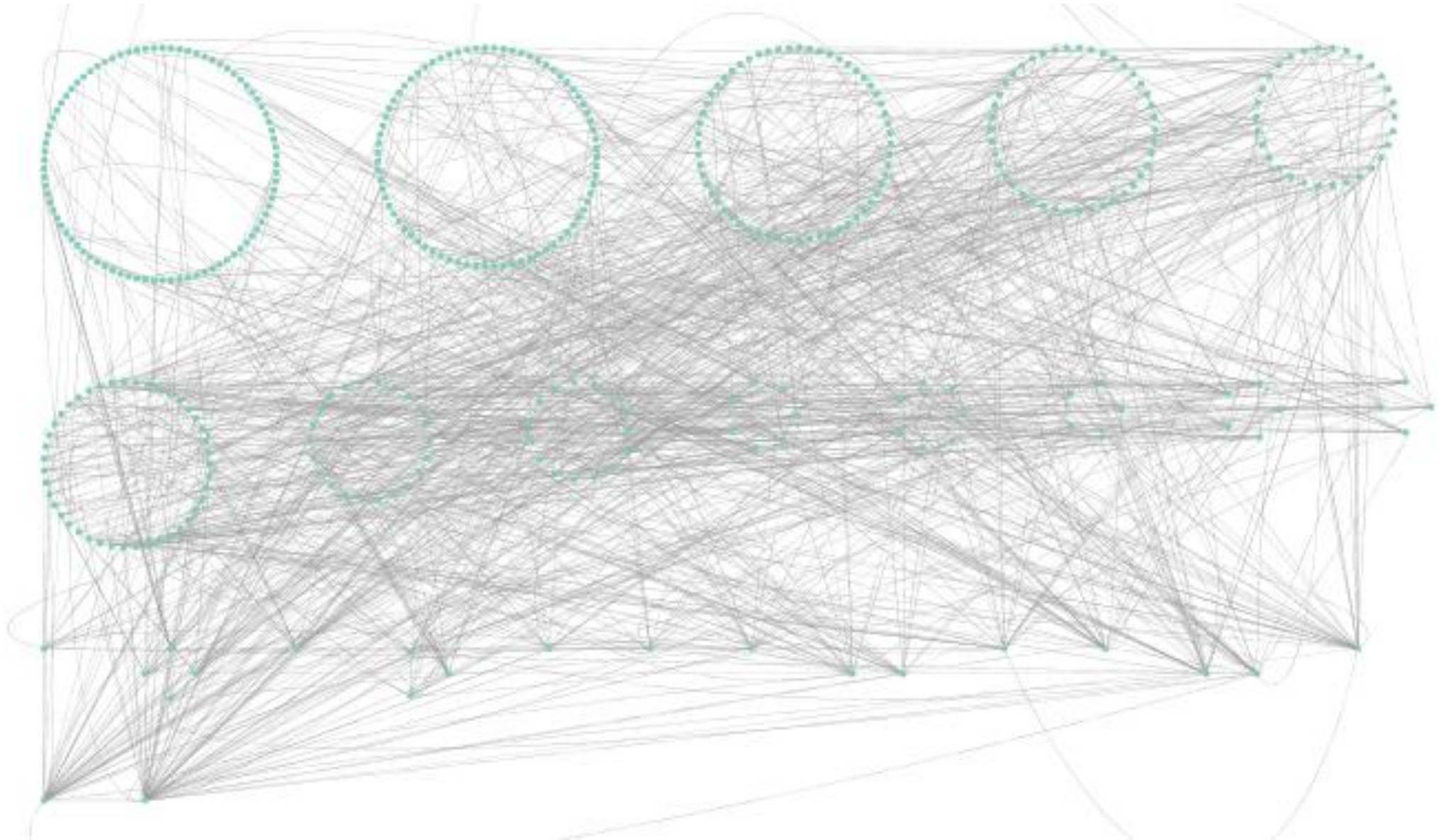
Advanced Options...

OK Cancel

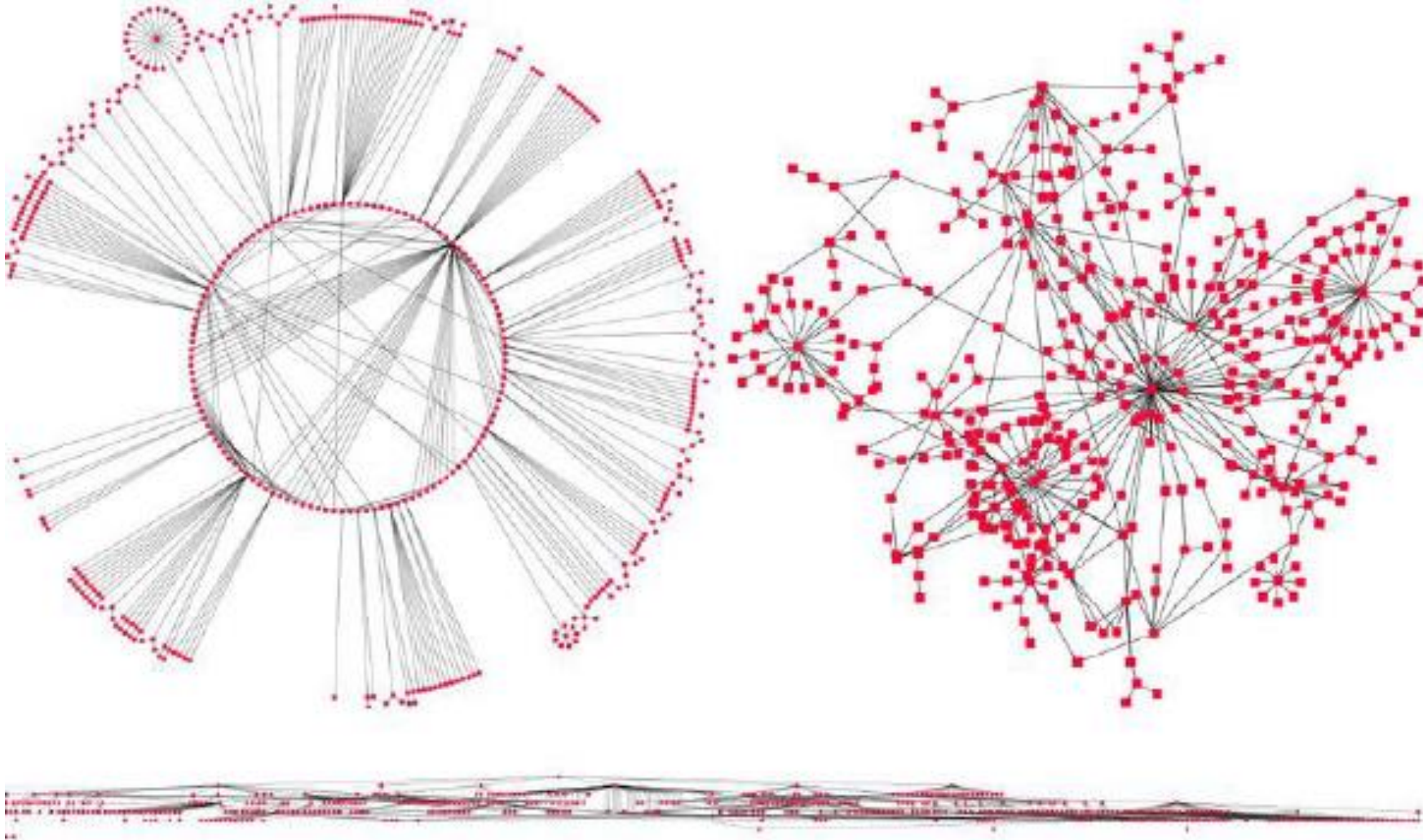
Layouts



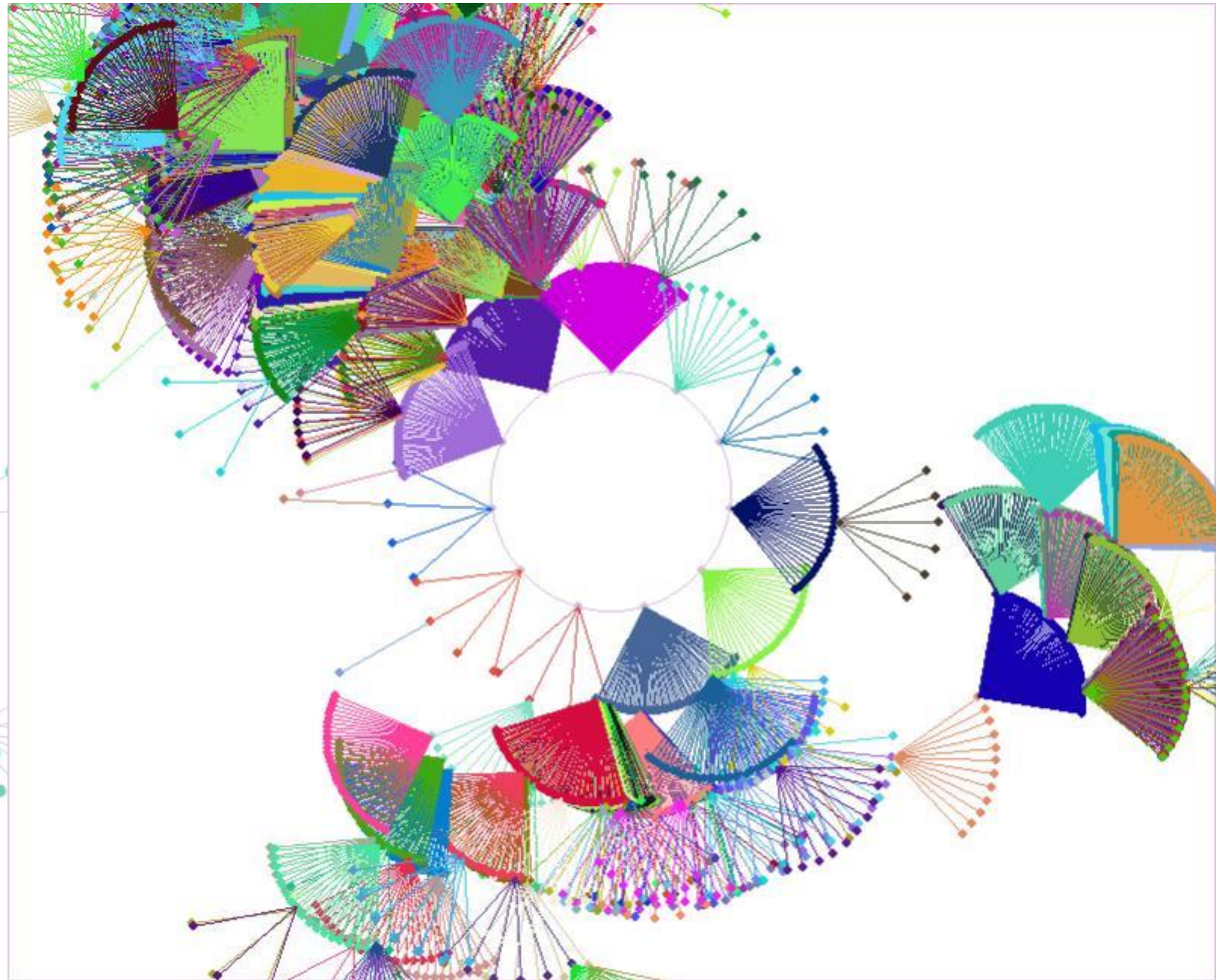
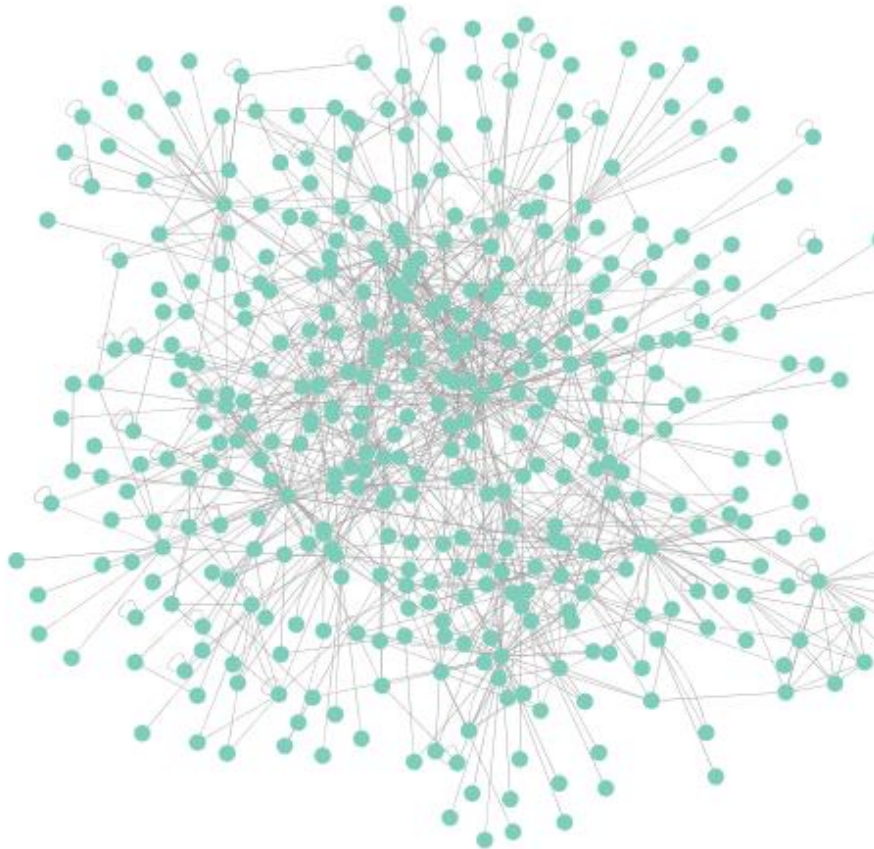
Layouts



Layouts




Layouts





Apps

<https://apps.cytoscape.org>

 Cytoscape App Store

Submit an App ▾

Search the App Store

Sign In

All Apps

Categories



[collections](#)
[data visualization](#)
[network generation](#)
[graph analysis](#)
[network analysis](#)
[online data import](#)
[integrated analysis](#)
[clustering](#)
[systems biology](#)
[utility](#)
[enrichment analysis](#)
[data integration](#)
[visualization](#)
[automation](#)
[layout](#)
[core app](#)
[pathway database](#)

Newest Releases

Get Started with the App Store »



OmniPath

3.0+

OmniPath: literature curated human signaling pathways



PathLinker

3.0+

Reconstructs signaling pathways from protein interaction networks



WikiPathways

3.0+

WikiPathways web service client and GPML file format importer



stringApp

3.0+

Import and augment Cytoscape networks from STRING



ModuLand 2.0

3.0+

Modularization method family offering modular hierarchies and



ClueGO

3.0+

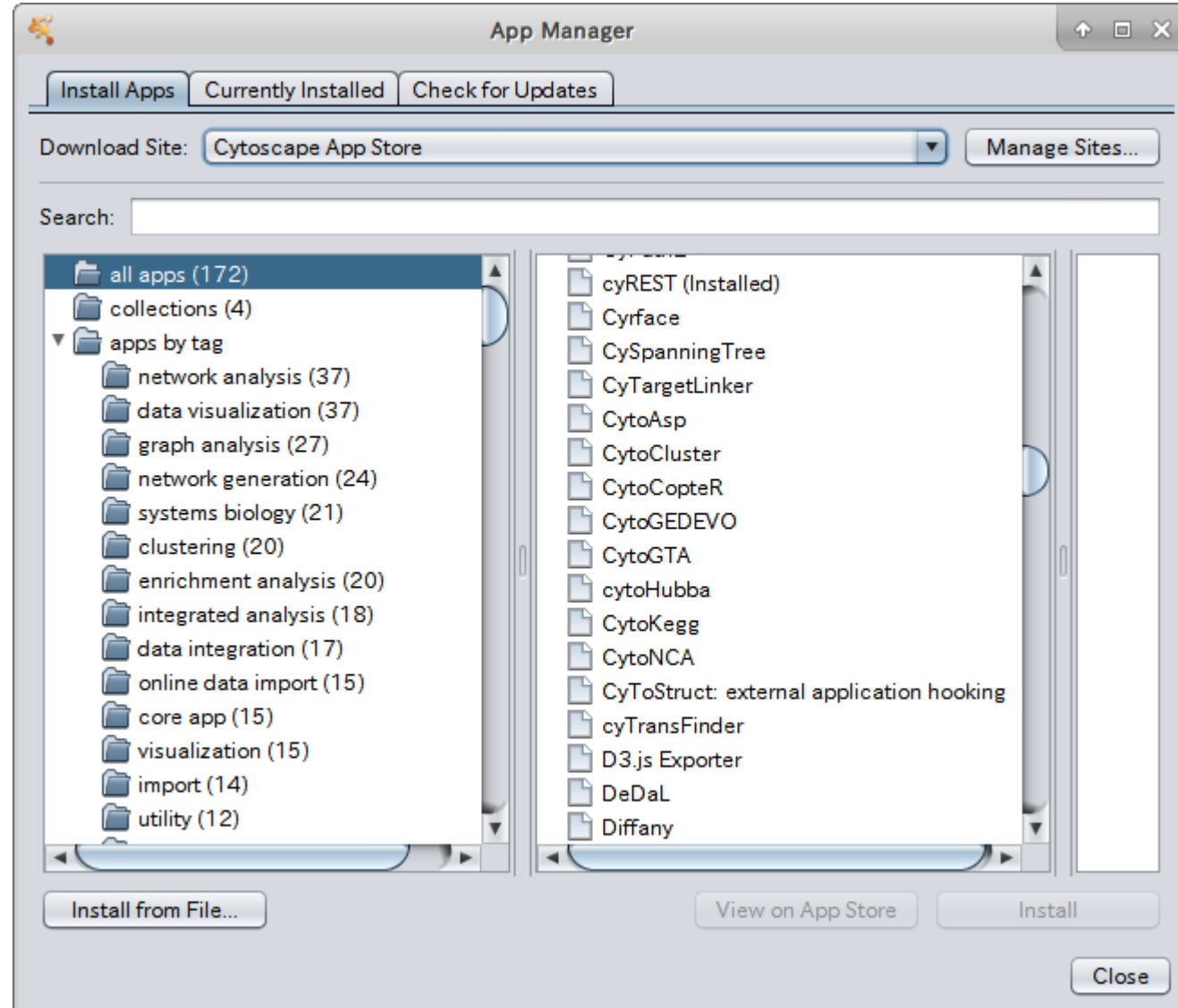
Creates and visualizes a functionally grouped network of

[more newest releases »](#)

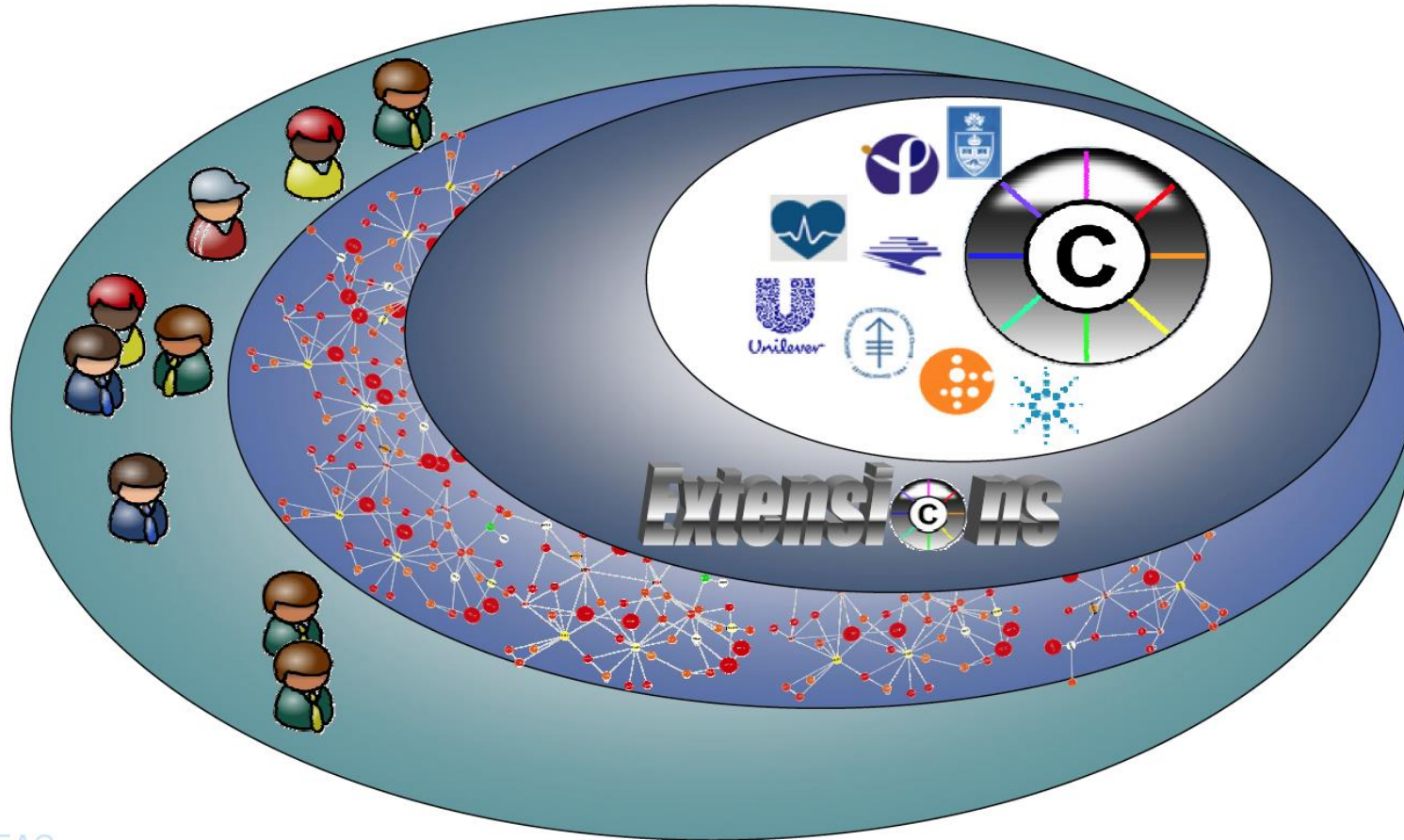
Top Downloaded Apps



Apps

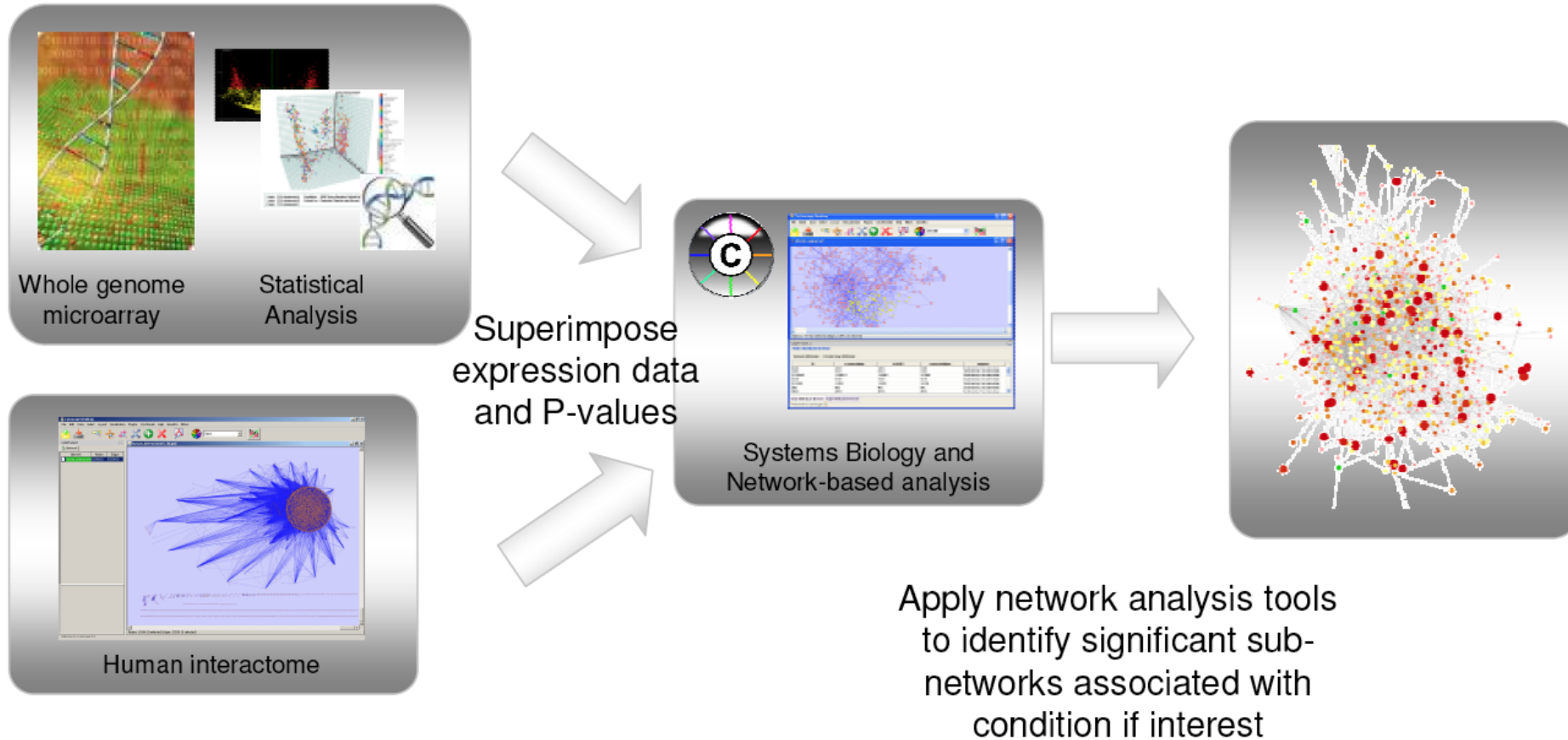


Cytoscape community



SEAC

Identify important subnetworks





Molecular Interaction and Network Resources for Cytoscape

- Import from web-services (NDEx, Pathway Commons)
- Sources: Mint, Reactome, IntAct, HPRD, Biogrid, SBCNY, MSKCC, BioGRID
- Pathway or Molecular interactions.
- Upload a network file (tab delimited, MITAB, SIF, CX).
- Build your own network!

Nature **437**, 1173-1178 (20 October 2005) | doi:10.1038/nature04209; Received 21 July 2005; Accepted 8 September 2005; Published online 28 September 2005

Towards a proteome-scale map of the human protein–protein interaction network

Jean-François Rual^{1,8}, Kavitha Venkatesan^{1,8}, Tong Hao¹, Tomoko Hirozane-Kishikawa¹, Amélie Dricot¹, Ning Li¹, Gabriel F. Berriz², Francis D. Gibbons², Matija Dreze^{1,3}, Nono Ayivi-Guedehoussou¹, Niels Klitgord¹, Christophe Simon¹, Mike Boxem¹, Stuart Milstein¹, Jennifer Rosenberg¹, Debra S. Goldberg², Lan V. Zhang², Sharyl L. Wong², Giovanni Franklin², Siming Li^{1,7}, Joanna S. Albala^{1,7}, Janghoo Lim⁴, Carlene Fraughton¹, Estelle Llamosas¹, Sebiha Cevik¹, Camille Bex¹, Philippe Lamesch^{1,3}, Robert S. Sikorski⁵, Jean Vandenhaute³, Huda Y. Zoghbi⁴, Alex Smolyar¹, Stephanie Bosak⁶, Reynaldo Sequerra⁶, Lynn Doucette-Stamm⁶, Michael E. Cusick¹, David E. Hill¹, Frederick P. Roth² & Marc Vidal¹

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4. Howard Hughes Medical Institute, and Departments of Pediatrics, Neurology, Neuroscience, and Molecular and Human Genetics, Baylor College of Medicine, One Baylor Plaza, Houston, Texas 77030, USA
5. Arcbay, Inc., 6 Whittier Place, Suite 7J, Boston, Massachusetts 01915, USA
6. Agencourt Bioscience Corporation, 500 Cummings Center, Suite 2450, Beverly, Massachusetts 01915, USA
7. †Present addresses: ArQule, Inc., 19 Presidential Way, Woburn, Massachusetts 01081, USA (S.L.); Departments of Cancer Biology, and Otolaryngology, Head and Neck Surgery, University of California Davis, 2521 Stockton Blvd, Suite 7200, Sacramento, California 95817, USA (J.S.A.)
8. *These authors contributed equally to this work

Correspondence to: David E. Hill¹Frederick P. Roth²Marc Vidal¹ Correspondence and requests for materials should be addressed to M.V. (Email: marc_vidal@dfci.harvard.edu), F.P.R. (Email: fritz_roth@hms.harvard.edu) or D.E.H. (Email: david_hill@dfci.harvard.edu).

Systematic mapping of protein–protein interactions, or 'interactome' mapping, was first performed in model organisms, starting with defined biological processes^{1,2} and then extending to the scale of the proteome^{3,4,5,6,7}. Although far from complete, such studies have revealed global topological and dynamic features of interactomes.



Enter pathway name or gene names



e.g. cell cycle, pcna xrcc2 xrcc3 rad50 rad51, P12004



NDEx

Browse

Search for networks, users, and groups



Featured Networks ▾

☐ Perform Search Term Expansion (Genes and Proteins only) ⓘ

Search Examples ▾

Featured Content



GeneHancer Associations

GeneHancer is a database of genome-wide enhancer-to-gene and promoter-to-gene associations, embedded in GeneCards.

NDEx, the Network Data Exchange:
A Resource for Network Analysis, Collaboration and Publication

Find Networks

Store and Share
Networks with
Collaborators

Use Networks in
Your Programs

Publish Your
Networks

NDEx is a Part of
Cytoscape



Try the new:

Integrated Query

Version 1.0

Analyze your gene set using networks in NDEx:
Pathway Enrichment - Protein Interactions - Drug, Disease, and Tissue Associations

Tweets by @NDExProject ⓘ



The NDEx Project

@NDExProject

@NDExProject is excited to be part of this long-awaited initiative! https://twitter.com/FAIRsharing_org/status/1217128681249951746



Jan 15, 2020



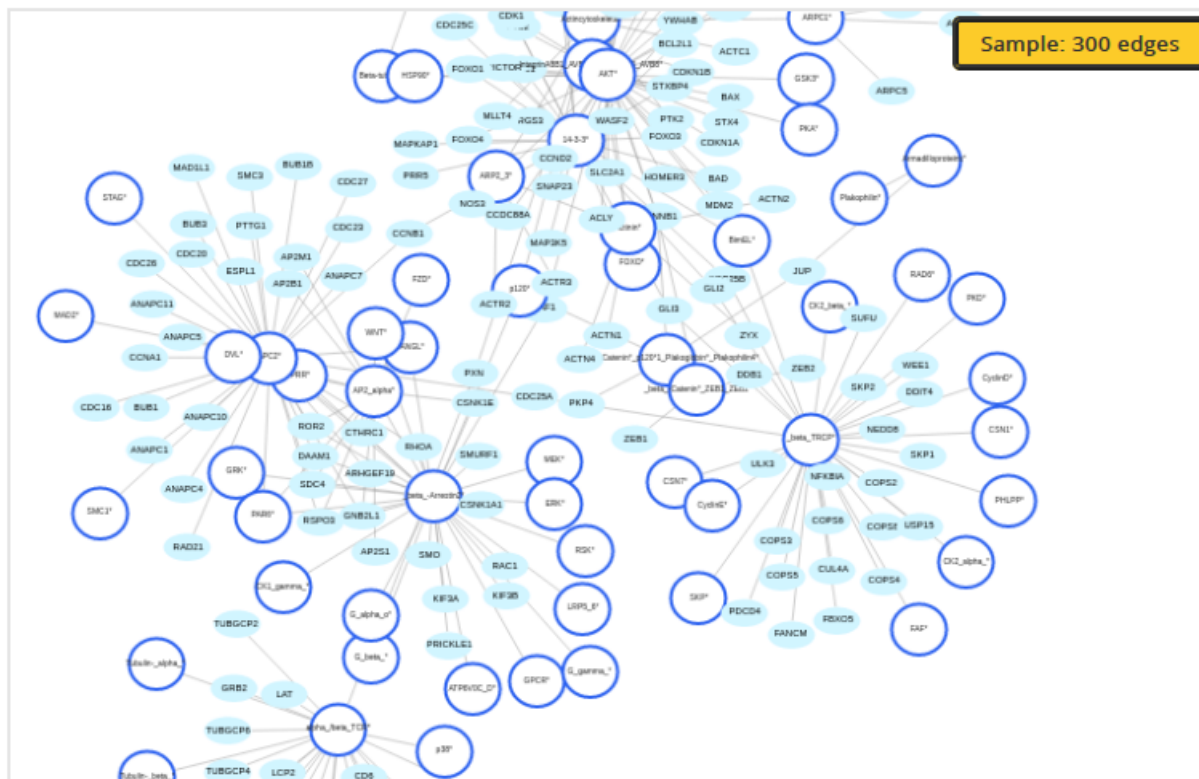
The NDEx Project Retweeted



Kleber Oliveira

@kleber_a_o

Our newest work is out! It resulted from a collaboration between @MACSIMaths and the Brazilian Federal Police. We examined over



Query Terms (i.e., AKT1 or WNT*)

Type: **i**

1-step neighborhood

Run Query

Network Info

Nodes/Edges

ACSN 2.0 Master - Protein-Protein Interactions

Nodes: 2540 Edges: 14090

PUBLIC ☒ Read Only

[Copy URL](#)

@context: [view namespaces](#)

Owner: [Atlas of Cancer Signalling Network](#)

Created: May 17, 2019 12:28:18 AM

Last Modified: Aug 12, 2019 11:27:51 PM

UUID: 49488c61-7832-11e9-848d-0ac135e8bacf

Format: Unknown

Description:

The [Atlas of Cancer Signalling Networks \(ACSN\)](#) is a web-based multi-scale resource of biological maps depicting molecular processes in cancer cell and tumor microenvironment.

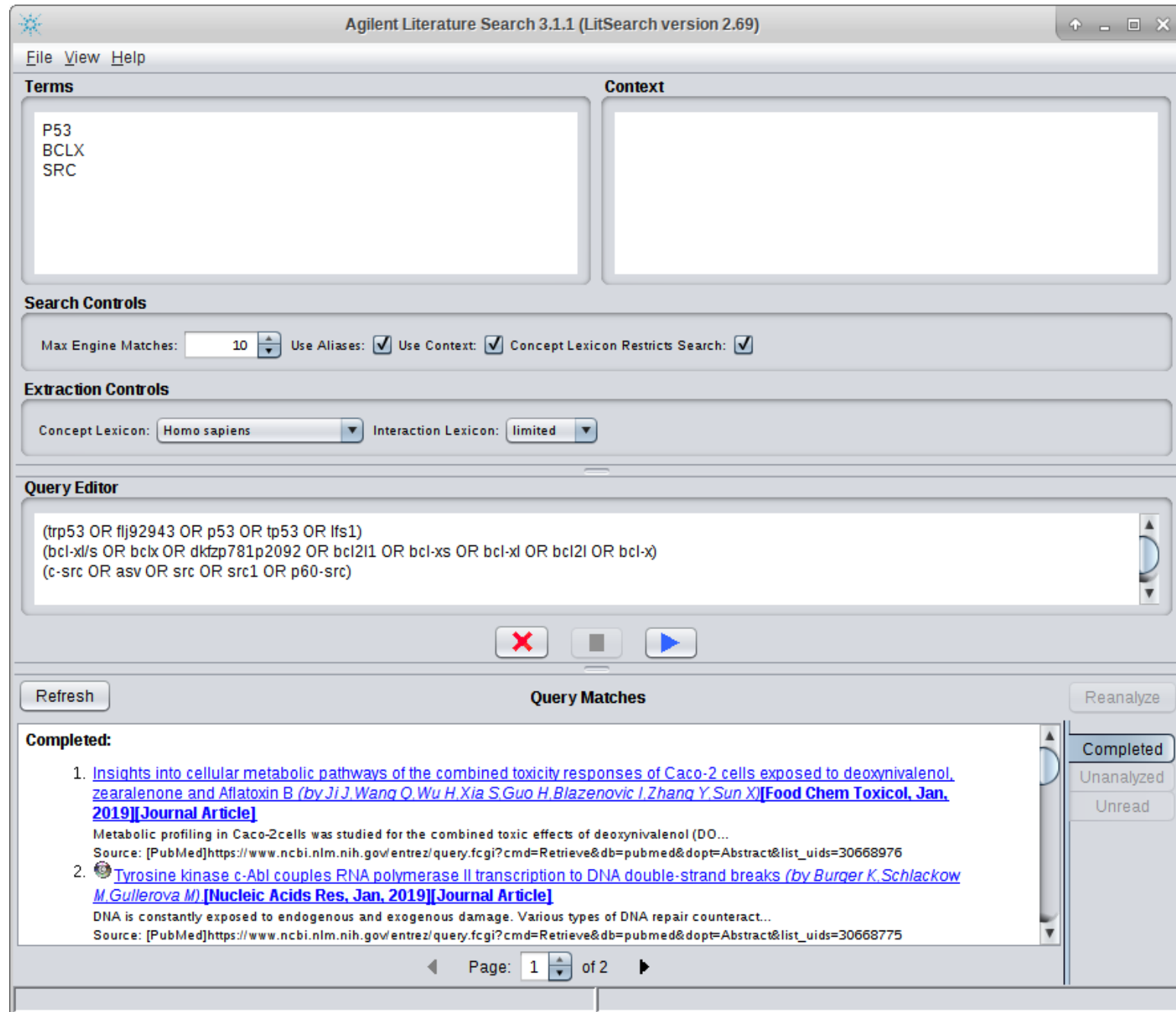
This is the Master Map containing all protein-protein interactions available in ACSN 2.0.

Rights Holder: Institut Curie



Table

Login





NCBI / PubMed API

A screenshot of the NCBI Developer Resources page. The page has a blue header with 'NCBI Resources' and 'How To' links. Below the header is a search bar with a 'PMC' dropdown and a 'Search' button. The main content area is titled 'Developer Resources' and contains text about using APIs and tools. It lists several APIs: NCBI E-utilities, File validation tools, OA web service, OAI-PMH, ID converter, FTP service, and Literature Citation Exporter.

NCBI Resources How To Sign in to NCBI

PMC US National Library of Medicine National Institutes of Health Advanced Journal list Help

About PMC For Publishers Related Resources

Developer Resources

We hope that developers make use of the APIs and tools that we have built for PMC.

In this page, we list the [APIs](#) and [public code repositories](#) related to PMC. We also present some of the [most common developer questions](#), and provide instructions and examples for addressing them in subsequent pages.

PMC APIs

The use of our APIs is entirely free, and doesn't require an API key, but we ask that you please:

- Do not make concurrent requests, even at off-peak times; and
- Include two parameters that help to identify your service or application to our servers:
 - *tool* should be the name of the application, as a string value with no internal spaces, and
 - *email* should be the e-mail address of the maintainer of the tool, and should be a valid e-mail address.

For the sake of readability, these parameters have been omitted from the *on-screen* display of the URLs in the example pages.

Here is a list of APIs that can be used to get data about PMC resources.

- [NCBI E-utilities](#),
- [File validation tools](#),
- [OA web service](#),
- [OAI-PMH](#),
- [ID converter](#),
- [FTP service](#),
- [Literature Citation Exporter](#).



Cytoscape API

Cytoscape 3.6.0 API

All Classes

Packages

org.cytoscape.app
org.cytoscape.app.event
org.cytoscape.app.swing
org.cytoscape.application
org.cytoscape.application.events

All Classes

AboutToRemoveEdgesEvent
AboutToRemoveEdgesListener
AboutToRemoveEdgeViewsEvent
AboutToRemoveEdgeViewsListener
AboutToRemoveNodesEvent
AboutToRemoveNodesListener
AboutToRemoveNodeViewsEvent
AboutToRemoveNodeViewsListener
AbstractBounded
AbstractCellEditor
AbstractConfigDirPropsReader
AbstractCyAction
AbstractCyActivator
AbstractCyApp
AbstractCyEdit
AbstractCyEvent
AbstractCyJob
AbstractCyNetworkReader
AbstractCyPayloadEvent
AbstractCySwingApp
AbstractEdgeViewTask
AbstractEdgeViewTaskFactory
AbstractFunction

Cytoscape 3.6.0 API

OVERVIEW PACKAGE CLASS USE TREE DEPRECATED INDEX HELP

PREV NEXT FRAMES NO FRAMES

Cytoscape Swing App API (swing-app-api) 3.6.0 API

This document represents the API specification for Cytoscape 3.0 using a Java Swing front-end.

See: Description

Packages

Package	Description
org.cytoscape.app	This is the Cytoscape App API, which supports development of Cytoscape 3.X apps in a manner similar to apps developed in Cytoscape 2.X.
org.cytoscape.app.event	
org.cytoscape.app.swing	This is the Cytoscape Swing App API, which supports development of Cytoscape 3.X apps in a manner similar to apps developed in Cytoscape 2.X and provides full access to the Swing specific services of the Cytoscape API in addition to all other services provided in Cytoscape App API.
org.cytoscape.application	This package provides Cytoscape version number, application-wide setting, basic access to current network, selected networks, views and rendering engines.
org.cytoscape.application.events	This package provides application events/listeners, including Cytoscape startup/shutdown, setCurrentNetwork/setCurrentNetworkView/ setSelectedNetwork.
org.cytoscape.application.swing	This package defines the various interfaces, abstract classes, and enums that represent the Cytoscape Swing Application API.
org.cytoscape.application.swing.events	This package defines the various events fired by the Cytoscape Swing Application API.
org.cytoscape.application.swing.search	



Cytoscape.js

Cytoscape.js: a graph theory library for visualisation and analysis

Max Franz; Christian T. Lopes; Gerardo Huck; Yue Dong; Onur Sumer; Gary D. Bader ✉

Bioinformatics (2015) 32 (2): 309-311.

DOI: <https://doi.org/10.1093/bioinformatics/btv557>

Published: 28 September 2015 **Article history** ▼

Views ▼ PDF Cite Share ▼ Tools ▼

Abstract

Summary: Cytoscape.js is an open-source JavaScript-based graph library. Its most common use case is as a visualization software component, so it can be used to render interactive graphs in a web browser. It also can be used in a headless manner, useful for graph operations on a server, such as Node.js.

Availability and implementation: Cytoscape.js is implemented in

JavaScript library for network visualization. Extendable with *Extensions* (not compatible with *Apps*).

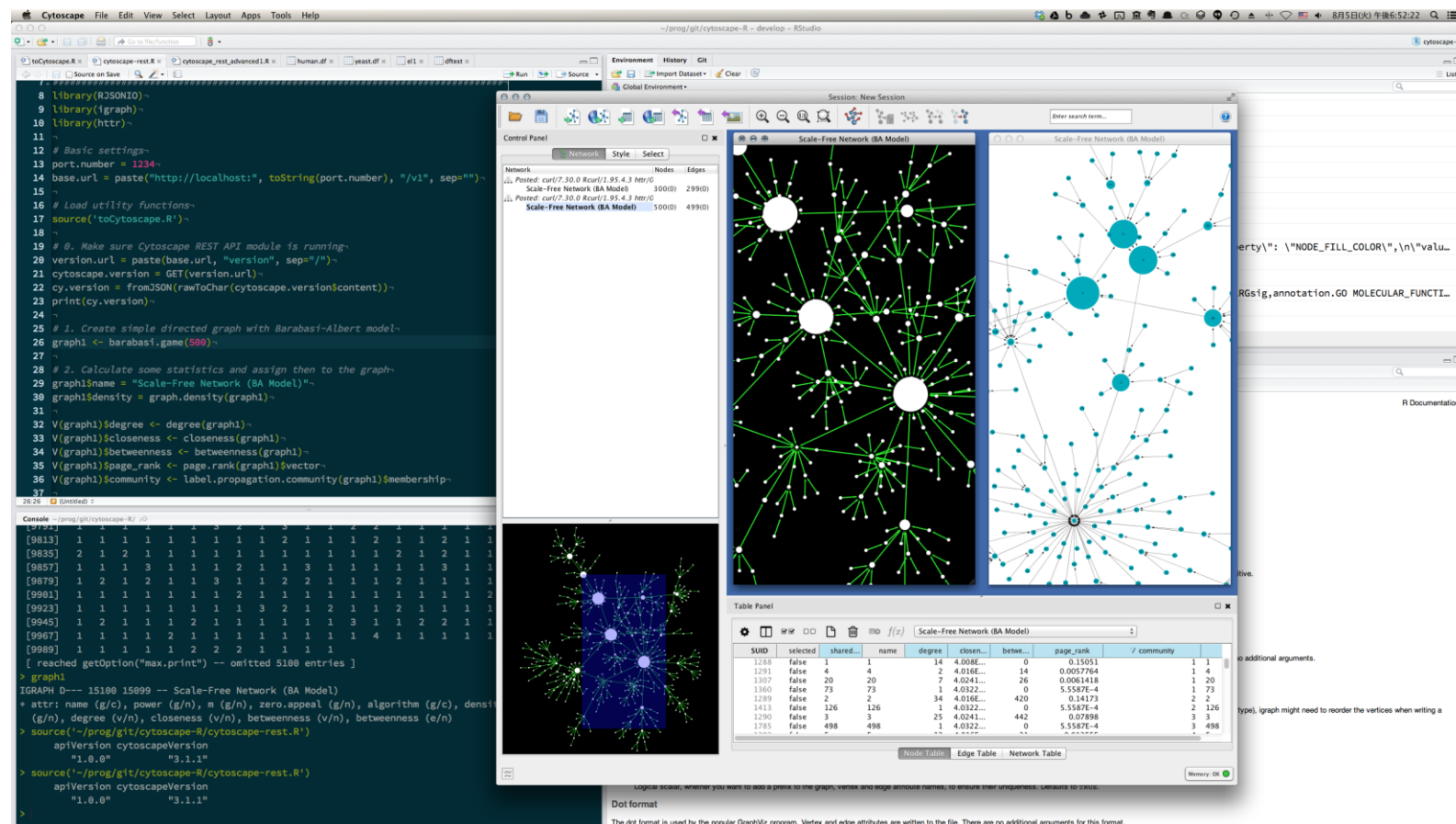
Long-term goal of closer integration between Cytoscape and Cytoscape.js.

You can export Cytoscape networks into Cytoscape.js and vice-versa.

<http://js.cytoscape.org/>



You can use the desktop Cytoscape app through a REST API, from R, Python, Jupyter notebooks, and other languages (it is a REST API after all...)



Ono, Keiichiro, et al. "[CyREST: Turbocharging Cytoscape Access for External Tools via a RESTful API](#)." F1000Research 4 (2015).

