

SysMO-DB and Virtual Liver SEEK: Sharing and Managing Systems Biology Experiments

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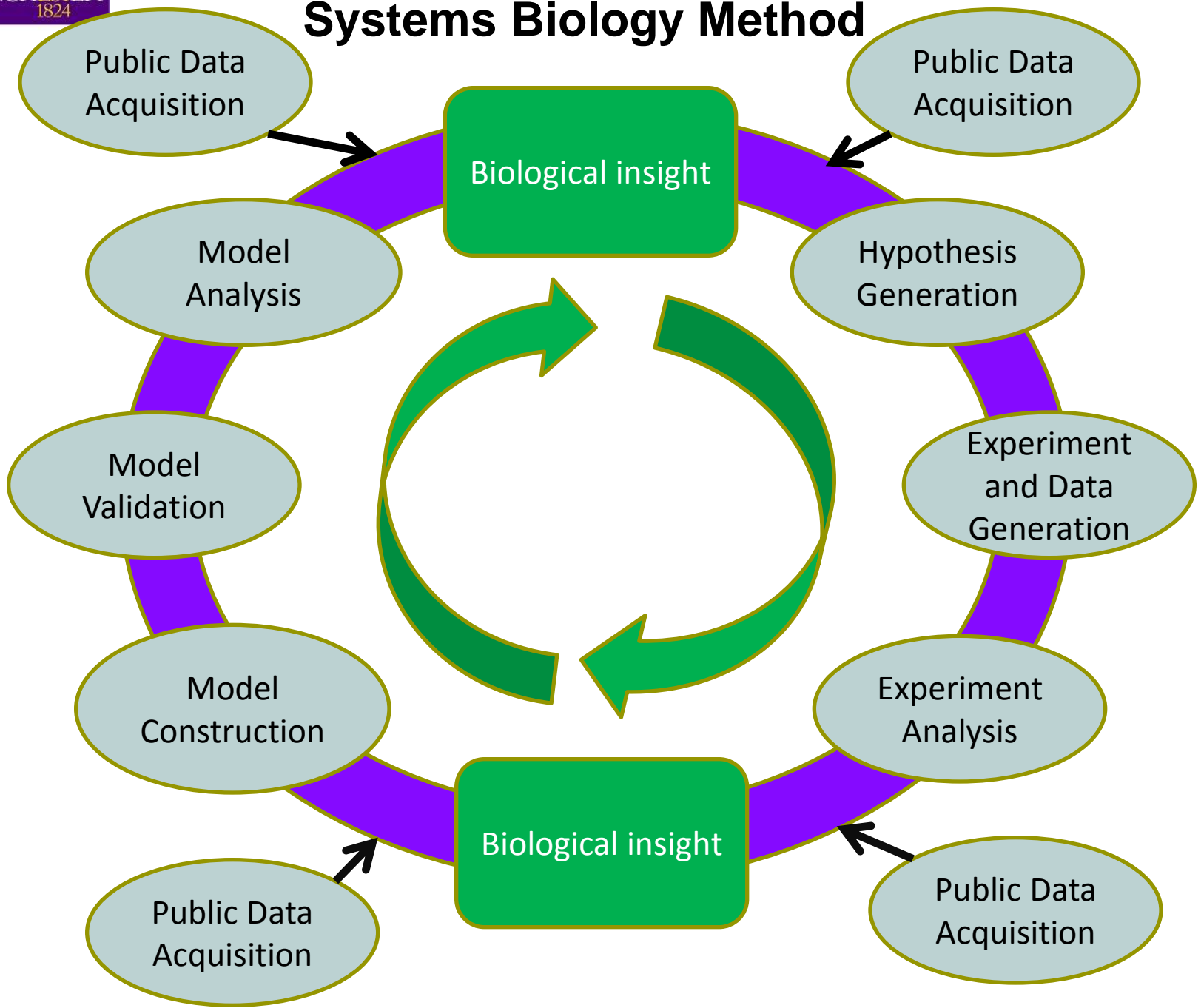
Martin Golebiewski

HITS

Systems Biology Method

Modelling

Experimental



SEEK: Systems Biology Data Sharing

- A platform for sharing Systems Biology data, models and protocols **in the context of systems biology experiments**
- Web based environment for sharing within a consortium, disseminating to the community, and exploring data and models (**an e-Laboratory**)
- **Standards compliant and standards promoting**
- Fitting in with laboratory practices

Assets Catalogue. Archive. Social Network. Sharing Space.



Find, share and exchange **Data, Models** and **Processes** within the SysMO Consortium.



People Projects Institutions Investigations Studies Assays Biosamples **Data** Models SOPs Publications Presentations Events Help

Provide Feedback

All

Admin

Yellow Pages:

People
Expertise
Projects
Institutions

I-S-A structure:

Linking
data,
models,
SOPs
Gateway to
BioPortal

Data:

Experimental data sets
and analysed results
Gateway to public data stores
– SABIO-RK, 'omics

Models:

Store
Simulate with JWS
Online
Annotate
Gateway to COPASI,
JWS Online, BioModels

Communication

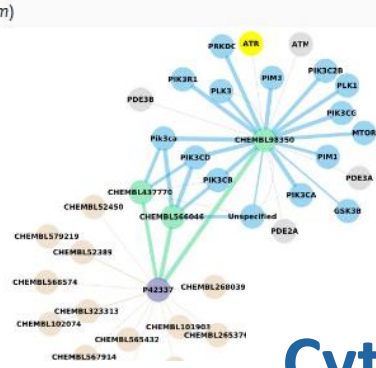
Publications
Community
Gateway to PubMed

Processes

Standard Operating Procedures
Computational workflows -
Taverna
Gateway to myExperiment

Versioning/Sharing

Exploring and Analysing: The SEEK as a Gateway



Cytoscape

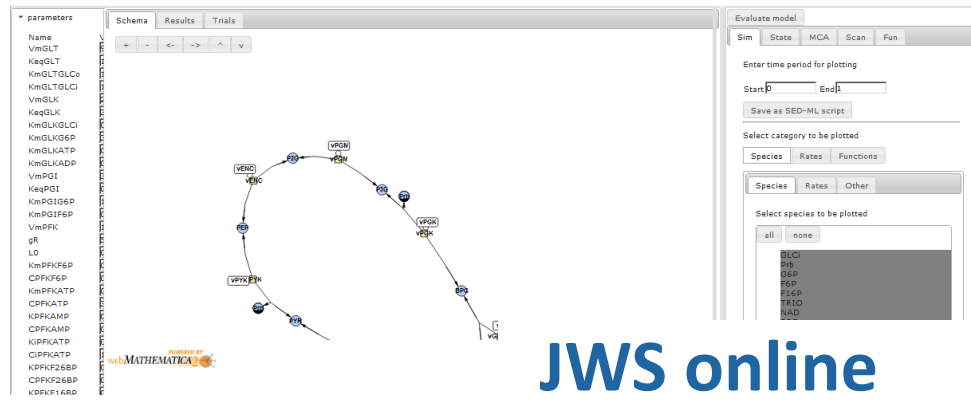
Visualise Model with Cytoscape Web

Download Model Simulate Model on JWS

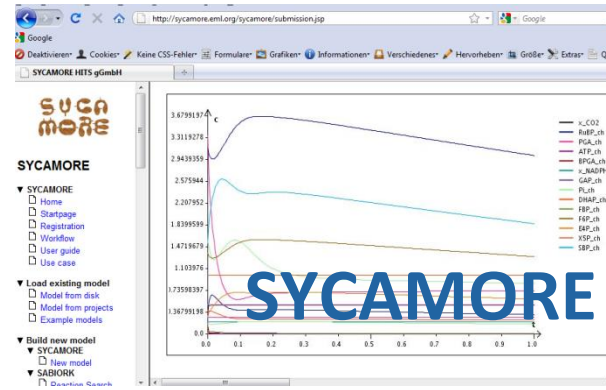
Simulate Model on Sycamore



Explore spreadsheets



JWS online



SYCAMORE



- Pan European collaboration
- 13 individual projects, >100 institutes
- Model dynamic molecular processes in microorganisms
- Pool research capacities and know-how
- Disseminate activities of the consortium

SEEK Communities



- German research network
- ~45 organisations, ~70 groups
- Multiscale representation of the liver
- Pool research capacities and know-how
- Disseminate activities of the consortium

Browse & Search by Scales

Find, share and exchange **Data**, **Models** and **Processes** within the Virtual Liver Network.



Yellow Pages

Experiments

Biosamples

Assets

Forums

Events

Help

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All

All

Go

Admin

New or upload

Data file

Go

Announcements

SEEK will be presented at ICSB 2012 and COMBINE / New SEEK version 4 months ago by Martin Golebiewski

Upload your data to SEEK as email attachment 5 months ago by Martin Golebiewski

New Virtual Liver SEEK with extended functionality 5 months ago by Martin Golebiewski

See all

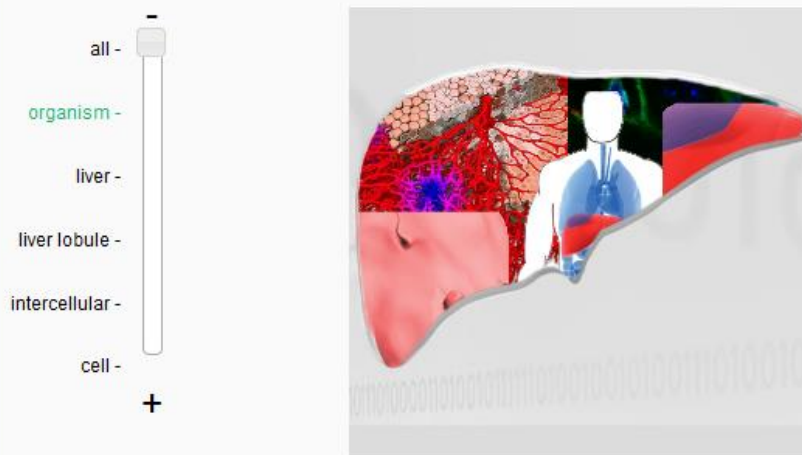
Favourites



Tags [show all]

CellNetAnalyzer (CNA) data management Flow Cytometry HepatoSys IL-6 Insulin

Search Items by Scale



Investigations (15+10)

Studies (11+13)

Assays (18+12)

Samples (29+5)

Specimens (33+9)

Data Files (263+131)

Models (59+13)

SOPs (33+1)

Publications (192)

Presentations (443+4)

Events (57+5)

Scales: Organism



Find, share and exchange **Data, Models** and **Processes** within the Virtual Liver Network.



Yellow Pages

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Biosamples

Assets

Forums

Events

Help

Provide Feedback

All

Organism

Go

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Favourites

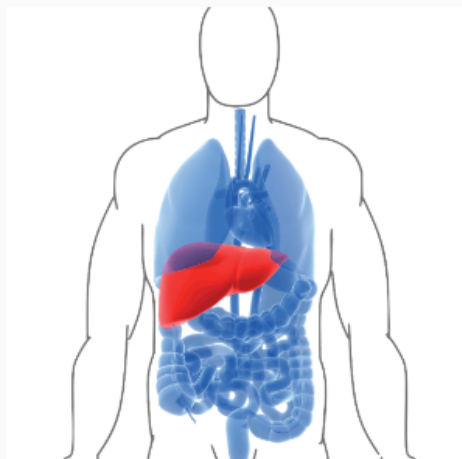


Tags [show all]

CellNetAnalyzer (CNA) data management Flow Cytometry HepatoSys IL-6 Insulin signalling Kinetic Modelling

Search Items by Scale

-
all -
organism -
liver -
liver lobule -
intercellular -
cell -
+



Specimens (17+1)

Data Files (8+3)

Models (2+1)

SOPs (2)

Publications (9)

Presentations (29)



Test mouse



Scales: Organ (Liver)

Find, share and exchange **Data**, **Models** and **Processes** within the Virtual Liver Network.



Yellow Pages

Experiments

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Forums

Events

Help

Provide Feedback

All

Liver

Go

Admin

New or upload

Data file

Go

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Favourites

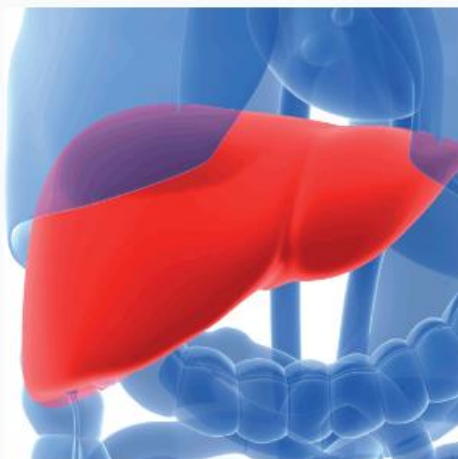


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all -
organism -
liver -
liver lobule -
intercellular -
cell -
+



Investigations (1+1)

Studies (2)

Assays (3)

Samples (12)

Specimens (8+1)

Data Files (31+14)

Models (2)

Publications (11)

Presentations (58)

Scales: Liver Lobule



Find, share and exchange **Data**, **Models** and **Processes** within the Virtual Liver Network.



Yellow Pages

Experiments

Biosamples

Assets

Forums

Events

Help

Provide Feedback

All

Liver Lobule

Go

Admin

New or upload

Data file

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Favourites

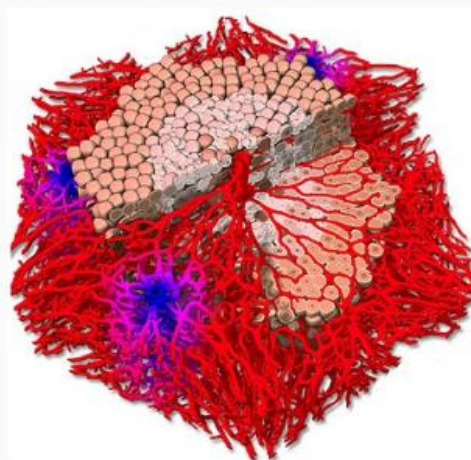


Tags [show all]

CellNetAnalyzer (CNA) data management Flow Cytometry HepatoSys IL-6 Insulin signalling Kinetic Modelling

Search Items by Scale

all -
organism -
liver -
liver lobule -
intercellular -
cell -
+



Investigations (1)

Samples (1)

Specimens (1)

Data Files (9+2)

Models (2+1)

SOPs (1)

Publications (5)

Presentations (46+1)

Scales: Intercellular



Find, share and exchange **Data, Models** and **Processes** within the Virtual Liver Network.



Yellow Pages

Experiments

Biosamples

Assets

Forums

Events

Help

Provide Feedback

All

Intercellular

Go

Admin

New or upload

Data file

Go



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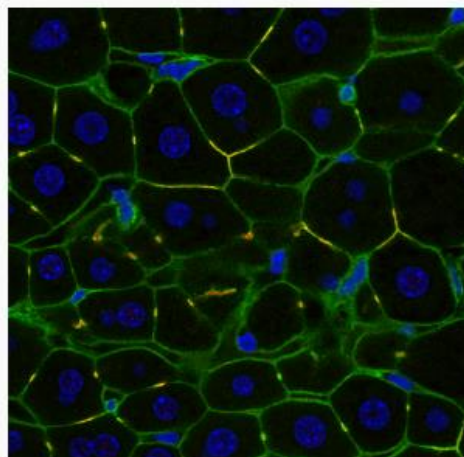
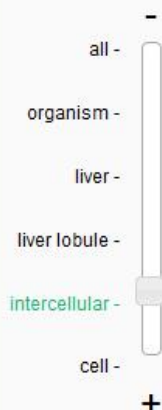
Favourites



Tags [show all]

CellNetAnalyzer (CNA) data management Flow Cytometry HepatoSys IL-6 Insulin signalling Kinetic Modelling

Search Items by Scale



Investigations (2+1)

Studies (1+1)

Assays (1)

Specimens (1)

Data Files (19)

Models (6+1)

SOPs (2)

Publications (13)

Presentations (65)



Scales: Cellular



Find, share and exchange **Data, Models** and **Processes** within the Virtual Liver Network.



Yellow Pages

Experiments

Biosamples

Assets

Forums

Events

Help

Provide Feedback

All

Cell

Go

Admin

New or upload

Data file

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See all

Favourites



Tags [show all]

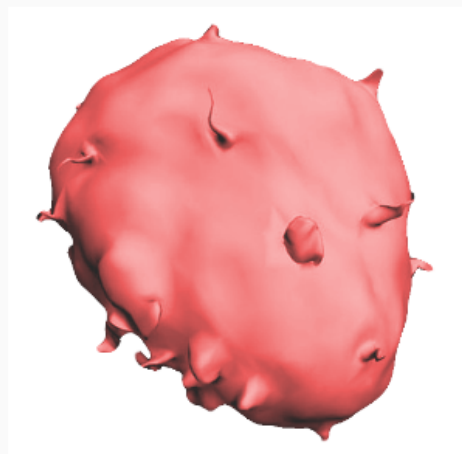
CellNetAnalyzer (CNA) data management Flow Cytometry

HepatoSys IL-6 Insulin signalling Kinetic Modelling

logical modelling metabolism

Search Items by Scale

all -
organism -
liver -
liver lobule -
intercellular -
cell -
+



Investigations (7+9)

Studies (6+10)

Assays (8+9)

Samples (10+1)

Specimens (6+2)

Data Files (69+56)

Models (54+7)

SOPs (11)

Publications (62)

Presentations (134+1)



Insulin signalling in primary hepatocytes



SEEK for Science

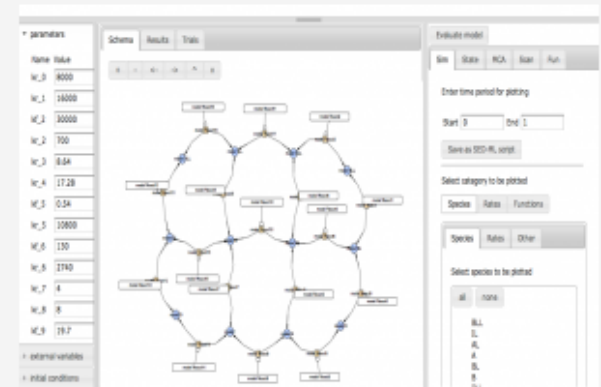
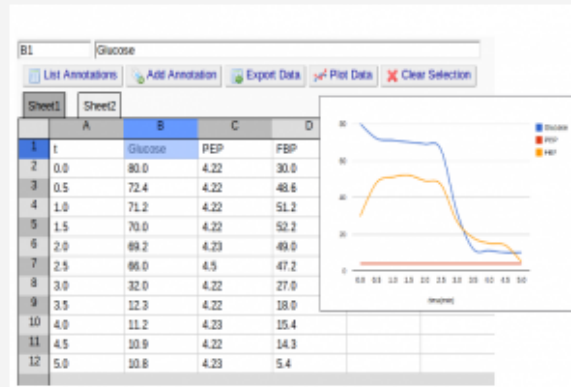
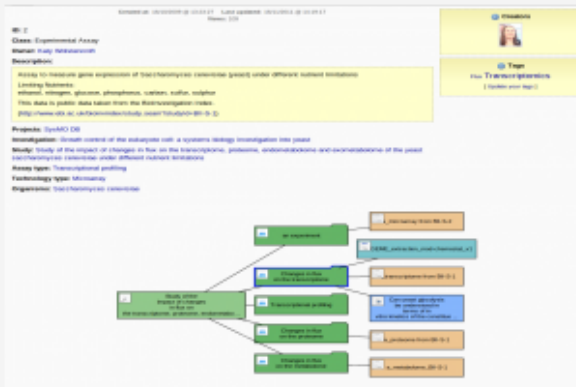
<http://www.seek4science.org>

For finding, sharing and exchanging Data, Models and Processes in Systems Biology.

Organise your experiments and data

Explore and annotate data

Simulate SBML models



SEEK has adopted an **ISATAB** style structure for organising experiments and data.

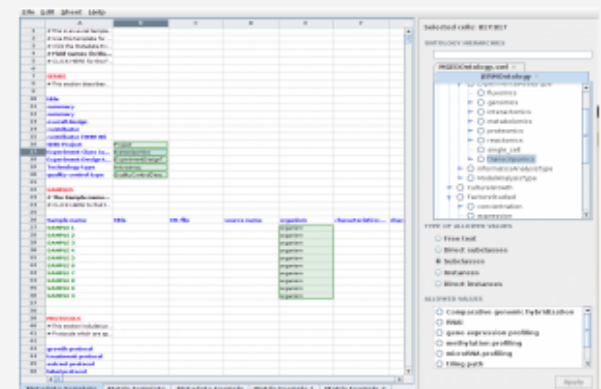
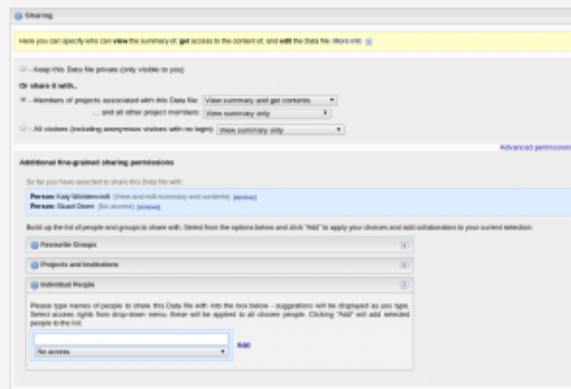
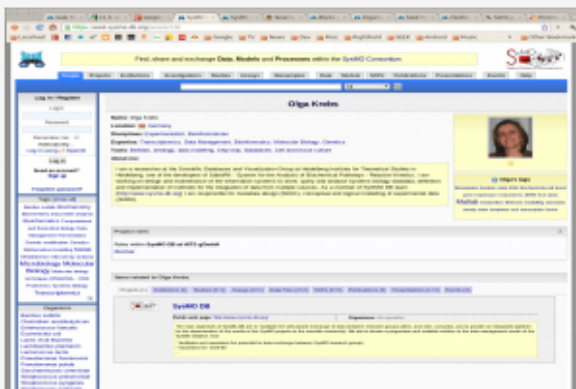
Excel spreadsheets can be explored and annotated without the need to download.

Most models that conforms to the **SBML** format can be simulated within SEEK.

Who's doing what, where?

Flexible sharing controls

Semantic spreadsheet templates



We recognise that people, and their knowledge, are important.

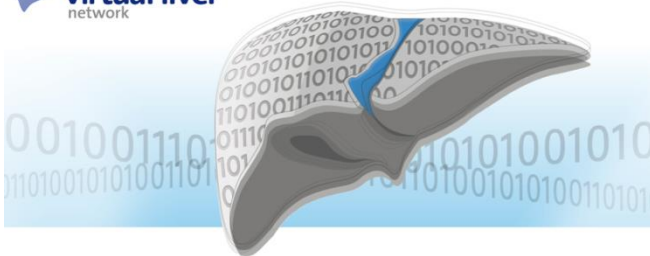
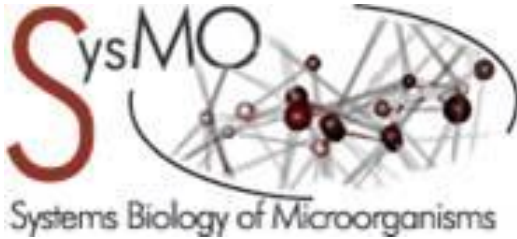
There is a lot of flexibility and control over who can see, download or edit your items.

Using **RightField** we are producing a wide collection of template files.



in Use

Developed for:



Adopted by:

SBCancer.



RosAge
Reactive oxygen species and
the dynamics of ageing

CISBIC

Consensus model
of Yeast Glycolysis



**THE SEEK BASIC PRINCIPLE: USE
WHAT IS AVAILABLE
NO REINVENTION
BUILD ON COMMUNITY ACTIVITIES**

**OBO**

Minimum Information Models

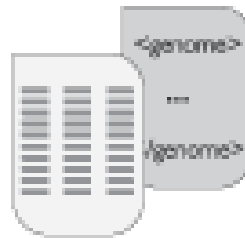
Formats, Ontologies,
Naming schemes and Controlled Vocabularies

What is the least amount of information required to:
Find, Interpret, Understand, Reuse

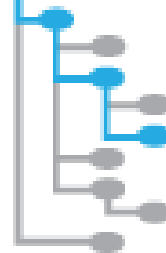
Realistic and pragmatic expectations for annotation
Ask for too much = people overwhelmed and provide less
Ask for too little = people will oblige

Transcriptomics
Proteomics
Metabolomics
30+ MIBBI

formats



terminologies



checklists



Matrix of Standards and Formats

biosharing

Minimal
Requirements

implements

Standard
Formats

add meaning

Ontologies

Data

Models

Simulation

Results



30+



?

MIAME
MIAPE

e.g. MAGE-TAB



SED-ML

NUML

isatab



JWS OneStop to Model Standards

SBML model format, SBGN schema, MIRIAM annotation

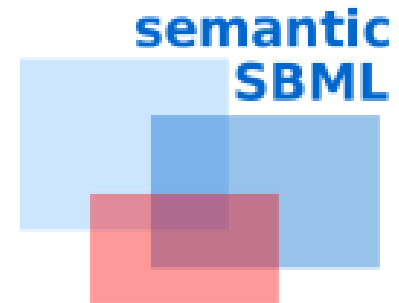
The screenshot displays the JWS OneStop software interface, which is used for creating and editing biological models. The interface is divided into several panels:

- Model name:** Contains a text field with the name "teusink" and a "Model name help" button.
- Reactions:** A list of reactions with their corresponding rate equations. For example, reaction v[2] is $\{1\}GLC_i + \{1\}Prb = \{1\}G6P$.
- Rate equations:** A large text area containing the full set of differential equations for the model, such as $v[1] = (VmGLT * (GLC_o - GLC_i / KeqGLT)) / (KmGLTGLC_o * (1 + GLC_o / KmGLTGLC_o + GLC_i / KmGLTGLC_i + (0.91 * GLC_o * GLC_i) / (KmGLTGLC_i * KmGLTGLC_o)))$.
- Annotations:** A panel on the right showing MIRIAM annotations assigned to the model. It includes a list of species (GLC_i, Prb, G6P, F6P, F16P, TRIO, NAD, BPG, P3G, P2G, PEP, PYR, ACE, NADH) and a table of annotations with checkboxes for selection.

At the bottom of the interface, there are buttons for "Initial values", "Parameter values", "Assignment rules", "Functions", and "Events".

SEEK integrated tool to:

- construct or modify models, save in SBML format
- get automated generation of SBGN schema
- annotate model (MIRIAM), implementing web services from semanticSBML.

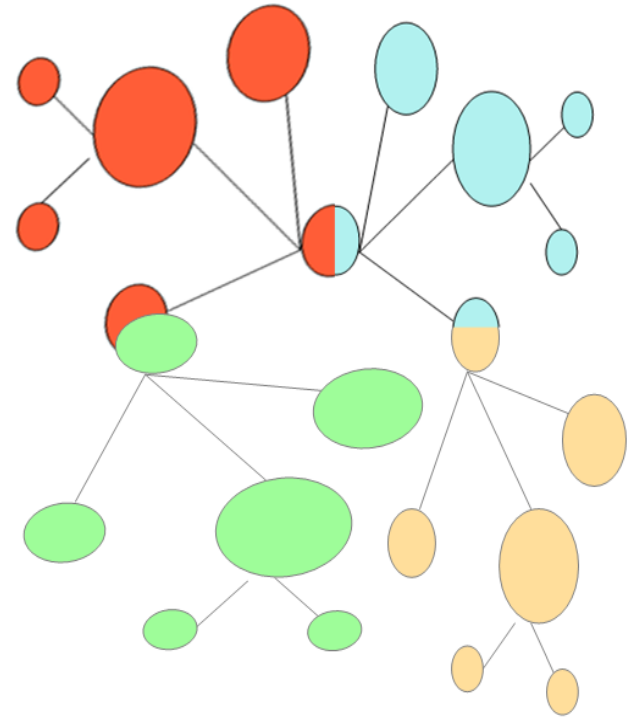
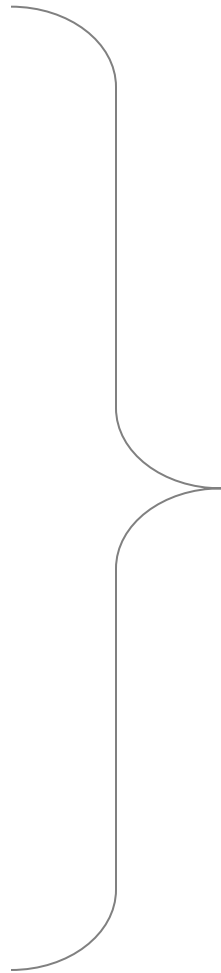
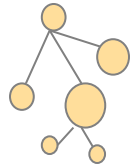
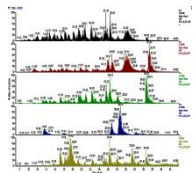
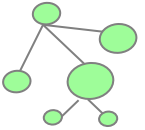
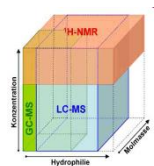
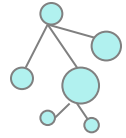
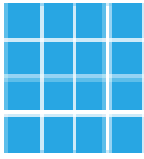
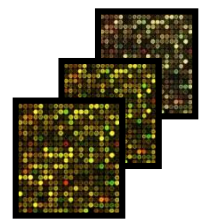
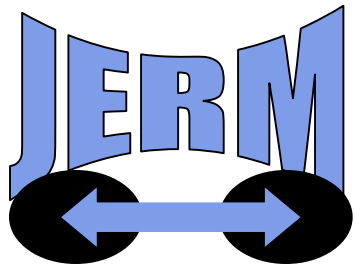


Omics data

standard
based
templates



RightField



RightField: Managing Vocabularies

RightField - C:\Users\katy\Dropbox\SysMo\Templates\Published\microarray_example_newformat.xls

File Edit Sheet Help

Excel workbook with marked-up cells

Selected parent term from the ontology

Methods for specifying ontology terms

Value Type and Property

Term lists for selected cells

Excel spreadsheet showing data entry fields and ontology terms:

	A	B	C
1	IDFI Investigation Description ...		
6			
7	# This section contains the top-l...		
8	Asset Title	Title	
9	Description		
10	Assay Title		
11	Experiment Class (AssayType)	Transcriptomics	
12	Experiment Description		
13	Experimental Design	DesignType	
14	Technology Type	microarray	
15	# Please create as many Experi...		
16	# describe the variables investig...		
17	Experimental Factor Name		
18	Experimental Factor Type	FactorType	
19			
20	# Quality Control Type examples...		
21	Quality Control Type	Quality Control Type	
22			
23	# Dates should be entered in the ...		
24	# it is recommended that you set...		
25	# to help avoid any unwanted ch...		
26	Public Release Date	YYYY-MM-DD	
27			
28	# Please list contact details in c...		
29	Person Last Name	name	
30	Person First Name		
31	Person SEEK ID	SEEKID	
32	SEEK Project	Project	
33	Person Email		
34	Person Phone		
35			

ONTOLOGY HIERARCHIES

MGEDOntology.owl x JERMOntology x

- Thing
 - Asset
 - Equipment
 - ExperimentalFactors
 - ExperimentDescription

VALUE TYPE AND PROPERTY

Subclasses

☒ Include a property

hasType

<http://www.mygrid.org.uk/ontology/JERMOntology#hasType>

ALLOWED VALUES

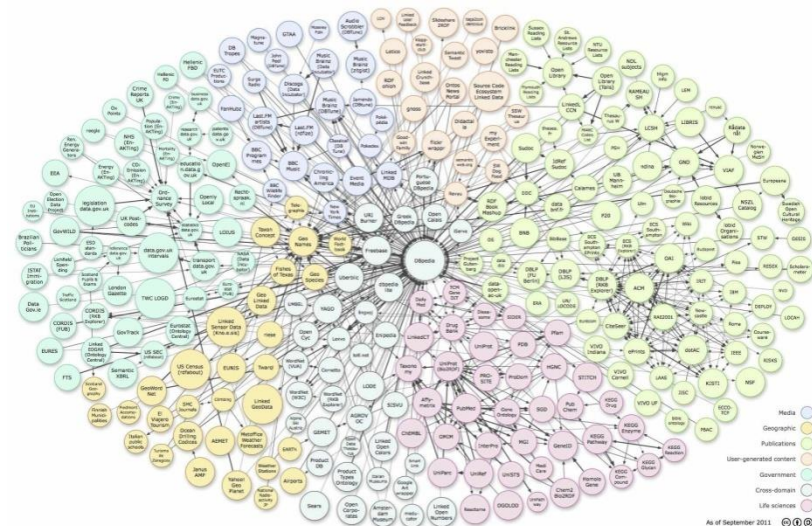
- ☐ gene expression profiling
- ☐ methylation profiling
- ☐ microRNA profiling

Apply

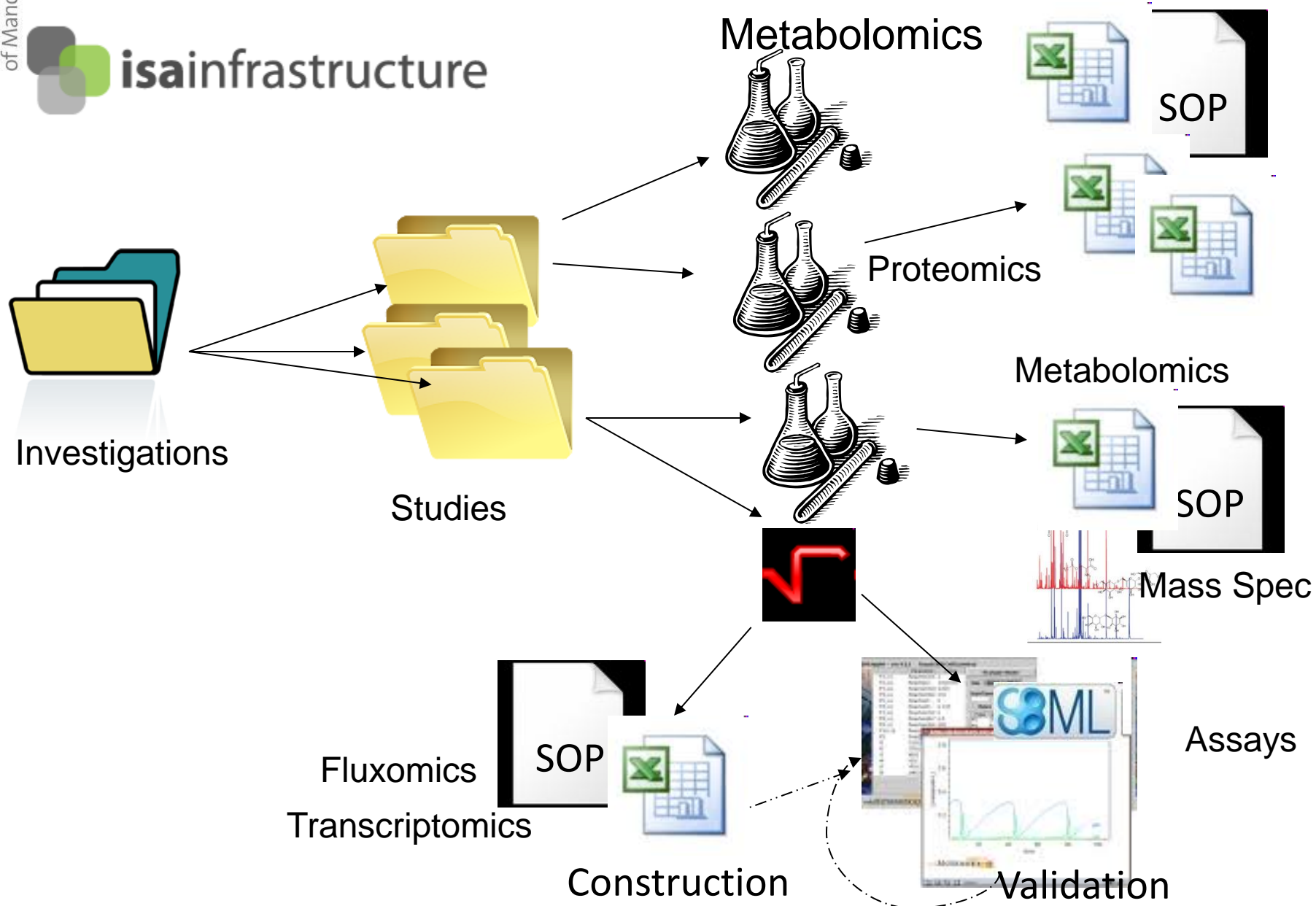
Semantic Linking and Querying

Extracting and storing metadata in RDF (Resource Description Framework), via RightField

- Better searching and querying
- New representations and visualisation of relationships between SEEK assets
- Linking SEEK data to the web of **Linked Data**

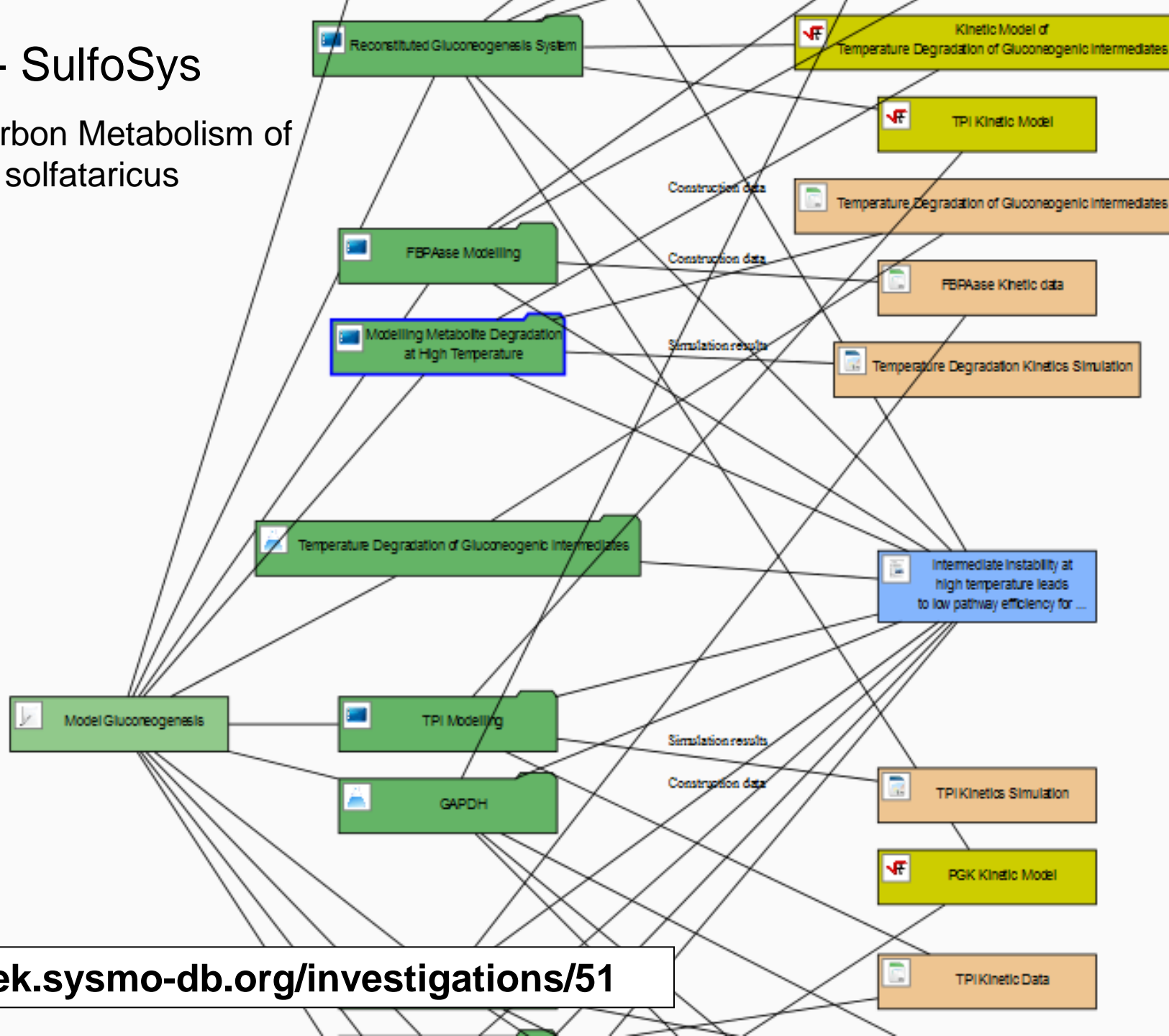


SEEK Links Data and Models



SysMO - SulfoSys

Central Carbon Metabolism of
Sulfolobus solfataricus



<https://seek.sysmo-db.org/investigations/51>

Incentives for Using SEEK

- Safe haven for Systems Biology assets
- Credit and attribution
 - Linking people to the data/models/sops etc
 - Linking assets to publications



- Publicity
 - SEEK records when data was uploaded/shared
 - Dissemination as well as internal project sharing

Incentives for using SEEK

- Exporting and linking
 - Supplementary materials store – persistent URIs
 - Linking publications and data
 - E.g. FEBS Journal
- Submission to public repositories
 - Silos not ideal for Systems Biology, but useful for meta-analysis

Why it works for us

- Off the shelf Systems Biology sharing environment
- Fits in with existing data and model management practices
- Incremental production with rapid prototypes and feedback from SEEK focus group (aka, the PALS)
- Publish and share within the consortium and beyond
- Scientists stay in control

Data management strategies for multinational large-scale systems biology projects

Wasco Wruck, Martin Peuker and Christian R.A. Regenbrecht

Submitted: 13th July 2012; Received (in revised form): 4th September 2012

Key Points

- Incentives to share data can be given by data citation credits (datacite).
- Open access to research data can be advanced via making sharing a condition of funding.
- Data management systems might be made attractive via alleviating and improving researchers' work, e.g. support for standard formats and publications.
- The reviewed systems proved useful for systems biology projects at least in dedicated environments, SysMO-SEEK out-of-the-box provides most useful features for large-scale systems biology projects.

Acknowledgements: SysMO-DB Team



Carole Goble



Stuart Owen



Katy Wolstencroft



Rob Haines



Niall Beard



Franco Du Preez



Jacky Snoep



Finn Bacall



Matt Horridge



Sergejs
Aleksejevs



Wolfgang Mueller



Olga Krebs



Quyen Nguyen



Lihua An



David Shockley



Martin
Golebiewski



Meik Bittkowski



Andreas Weidemann