BioModels

Sharing and re-using computational models of biological processes

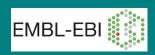
Nicolas Le Novère, Babraham Institute, EMBL-EBI





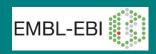
The world before BioModels

- List of models on webpages. Generally a link to the model in some format
- Main repositories were ModelsDb (neuroscience), DOQCS (neuroscience), the CellML and the SBML repositories. Plus a handful of personal web pages
- 100% of the models in the SBML repository were wrong. Wrong structure, wrong parametrisation, or did not produce the correct result
- It was generally not possible to test the other repositories because of the lack of tools



What is BioModels?

- Store and serve quantitative models of biological interest
- Models described in the peer-reviewed scientific literature + models automatically generated from pathway resources
- Models are curated: computer software check the syntax, while human curators check the semantics
- Models are simulated to ensure they provide the expected results
- Model components are annotated, to improve identification and retrieval
- Models are accepted in several formats, and served in several others



What can-we do with BioModels Database?

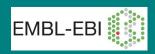
- Search and retrieve relevant quantitative or qualitative models
- Explore the variables and the structure of the model
- Run timecourse simulations of the model
- Download the model in many formats
- Do all the above via Web Services
- Learn about significant models using the Model of the Month



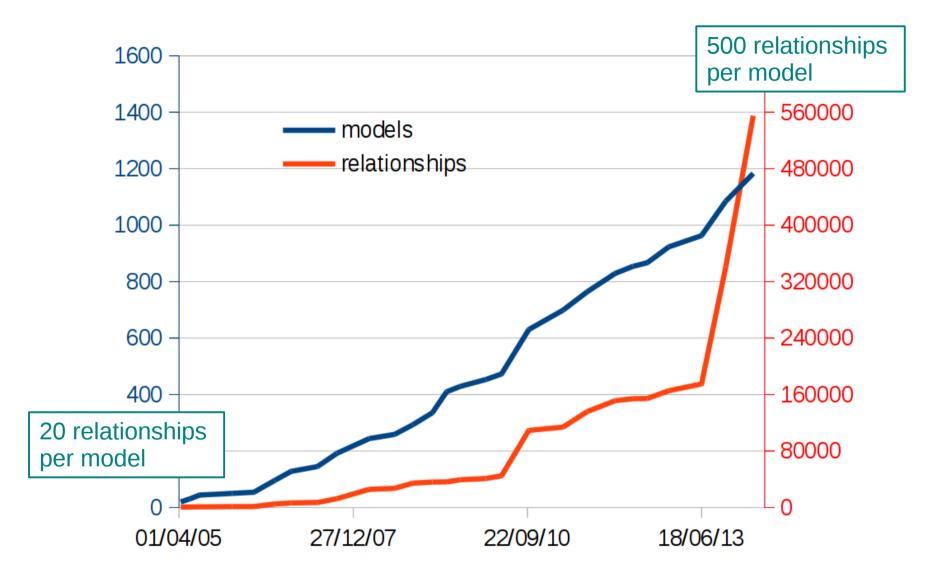
Where do models come from?

- From authors before grant applications or publications
- Over 300 scientific journals advise submission to BioModels Db:
 - Molecular Systems Biology
 - Public Library of Science journals
 - BioMedCentral journals
 - Royal Society of Chemistry journals
- Various people curated models out of interest.
- Submitted by curators
 - imported from other repositories (DOQCS, CellML)
 - reimplemented from literature



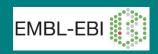


Computational models on the rise

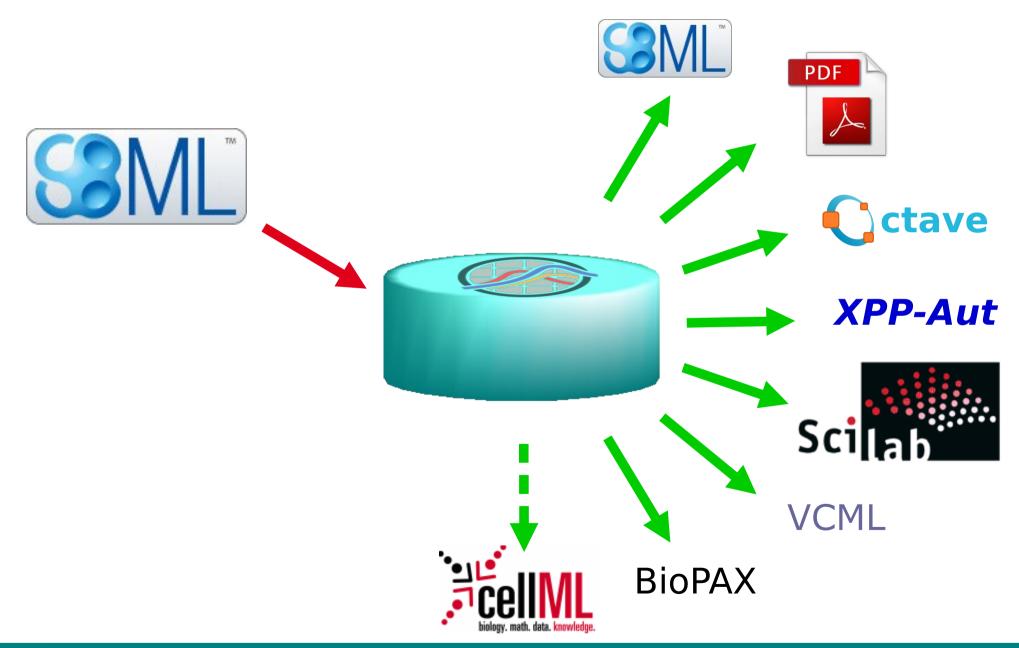


BioModels Database growth (published models branch) since its creation





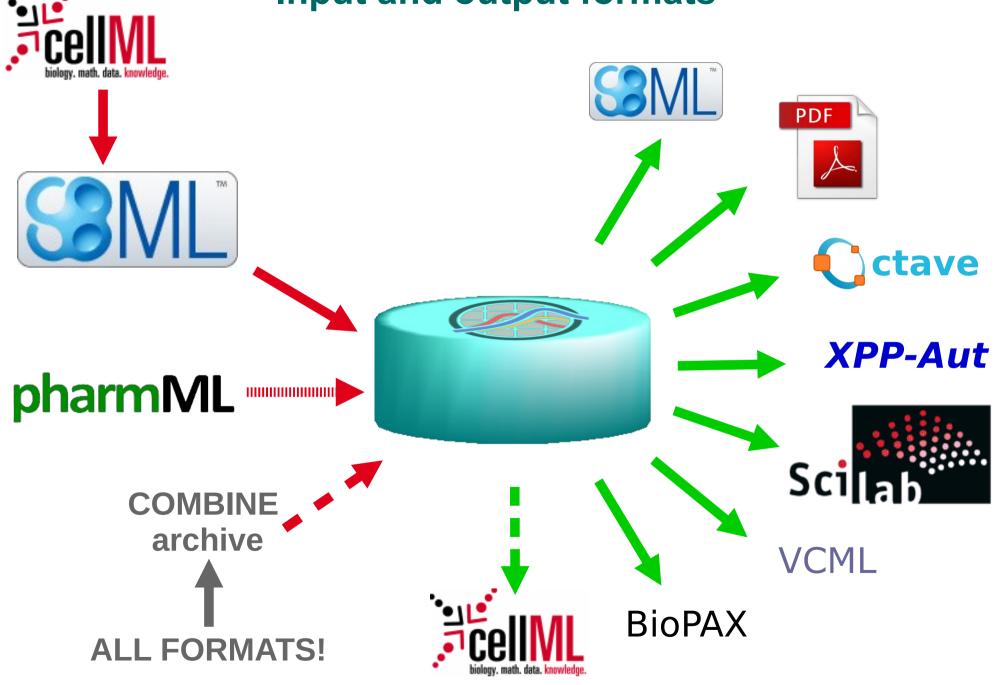
Input and output formats







Input and output formats



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 - 1.3 Online converters service
 - 1.4 Development
 - 1.5 Getting Help and Support
- 2 Other Converters

Systems Biology Format Converter

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The System Biology Format Converter

The System Biology Format Converter (SBFC) aims is to provide a generic framework that potentially allows any conversion between two formats. Interoperability between formats is a recurring issue in Systems Biology. Although there are various tools available to convert models from one format to another, most of them have been independently developed and cannot easily be combined, specially to provide support for more formats. The framework is written in Java and can be used as a standalone executable. This is a collaborative project and we hope that developers will provide support for more formats by creating new modules. SBFC allows anyone to easily add new converters and to integrate existing converters with a minimum of changes. We will also allow to combine several existing converters.

The SBFC framework currently supports conversion from SBML 🗗 to BioPAX 🗗 Level 2 and Level3, XPP 🗗, Octave 🗗 and Dot 🗗.

Several others converters are in development:

- SBGN & related:
 - SBML to SBGN-ML
 - · SBGN-ML to an SBGN enriched graphical representation designed for use in a web browser

The main Sourceforge page for the SBFC project is http://sourceforge.net/projects/sbfc/

Converters

Here is a small description of all the conversions currently supported. You can access to a more detailed page by clicking on a converter name.







Step 1 : Choose your input format

The input format you choose should match the model(s) you will submit, otherwise you will get an error. We strongly advise you validate your model(s) before the conversion

At the moment only SBML files can be selected as input.

Model format : SBML ▼

Step 2: Choose your output format

The output format you choose will determine the converter to use, if the format you want doesn't appear here it means the converter is not implemented yet

Output format : Select the output format Select the output format BioPax Level 2 BioPax level 3 Octave

Step 3 : Enter XPP

SBML L3V1

SBML_L2V4 SBML_L2V1 SBML_L1V2 et a URL to your results (optional)





BioModels is not limited to biochemistry!

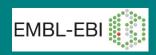
Is it about bio[logy|chemistry|medicine]?

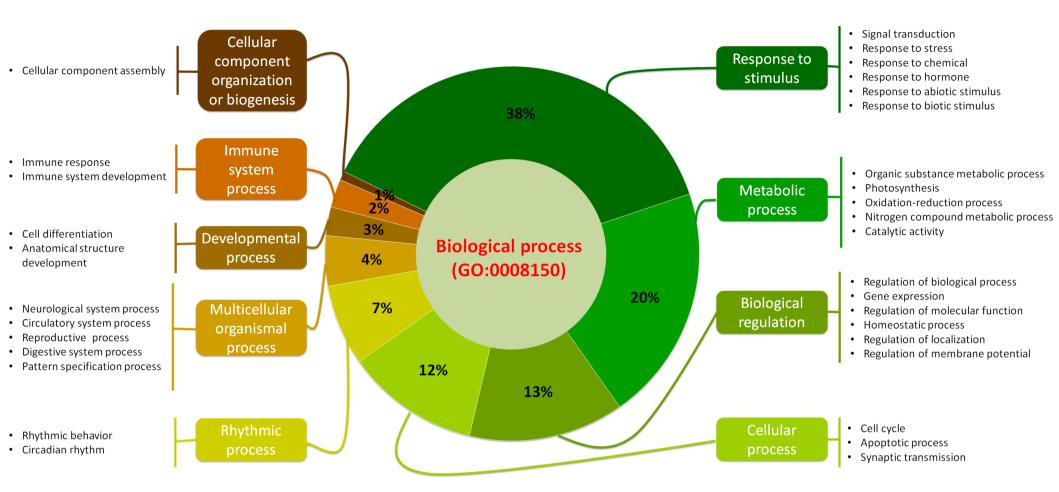
Is it a computational model?

Is it published?

Then it should be in BioModels



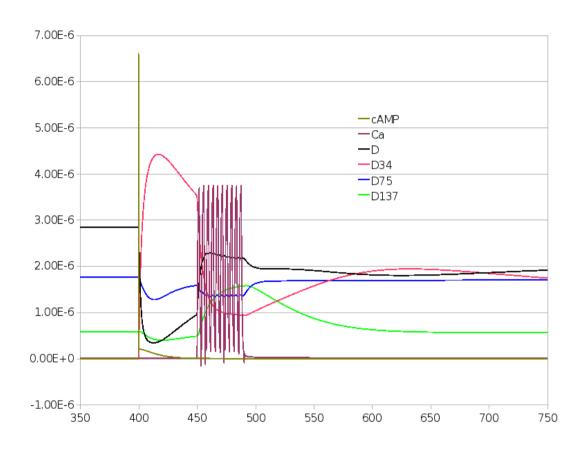








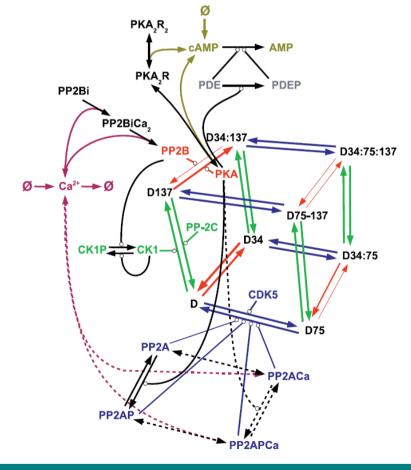
Biochemical models



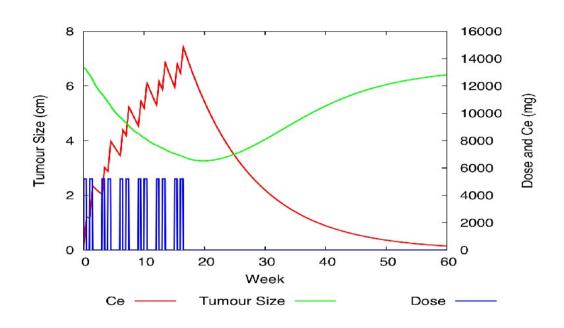
reaction: $v_{\rm on} = k_{\rm on} \times [{\rm D}] \times [{\rm CDK5}] \times {\rm Vol}$

Fernandez et al. DARPP-32 is a robust integrator of dopamine and glutamate signals *PLoS Comput Biol* (2006) 2: e176.





Pharmacometrics models



Tham et al (2008) A pharmacodynamic model for the time course of tumor shrinkage by gemcitabine + carboplatin in non-small cell lung cancer patients.

Clin Cancer Res. 2008 14(13): 4213-8.



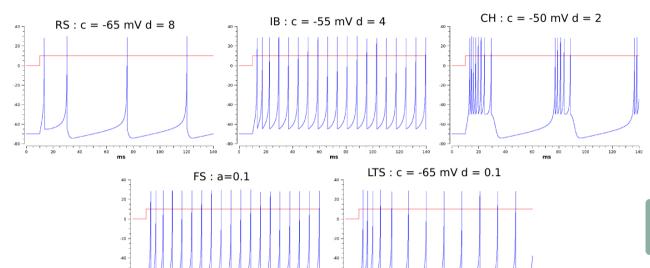
$$\frac{dSize}{dt} = (Rate_{in} \times Effect - K_{over} \times Size) \times Size$$

assignment rule:

$$Effect = 1 - \frac{E_{max} \times Ce}{Amt_{50} + Ce}$$



Neuroscience models



Izhikevich EM. Simple model of spiking neurons. IEEE Trans Neural Netw (2003) 14(6):1569-1572.

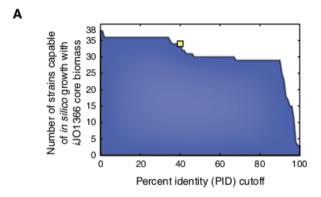


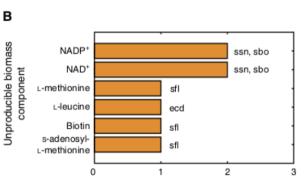
rate rule:

$$\frac{dv}{dt} = 0.04^2 + 5 \times V + 140 - U + i$$

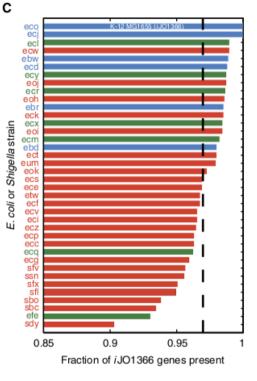
event:

Flux Balance Analysis models





Number of strains



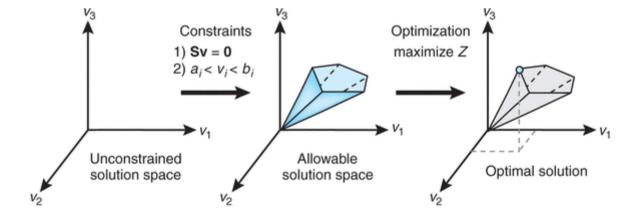
Orth et al. A comprehensive genomescale reconstruction of Escherichia coli metabolism - 2011

Mol Syst Biol (2011);7:535

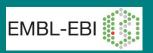


reaction:

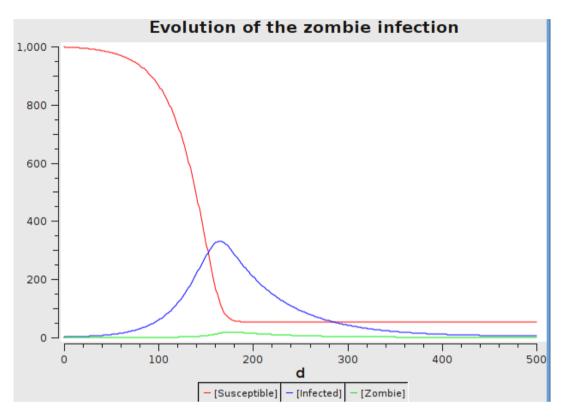
$$v_i = flux_i$$







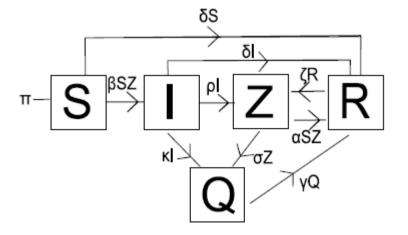
Epidemiology models





Munz P et al. When zombies attack!: Mathematical modelling of an outbreak of zombie infection. in "Infectious Disease Modelling Research Progress", (2009)133-150





BioModels Db does not belong to anyone!

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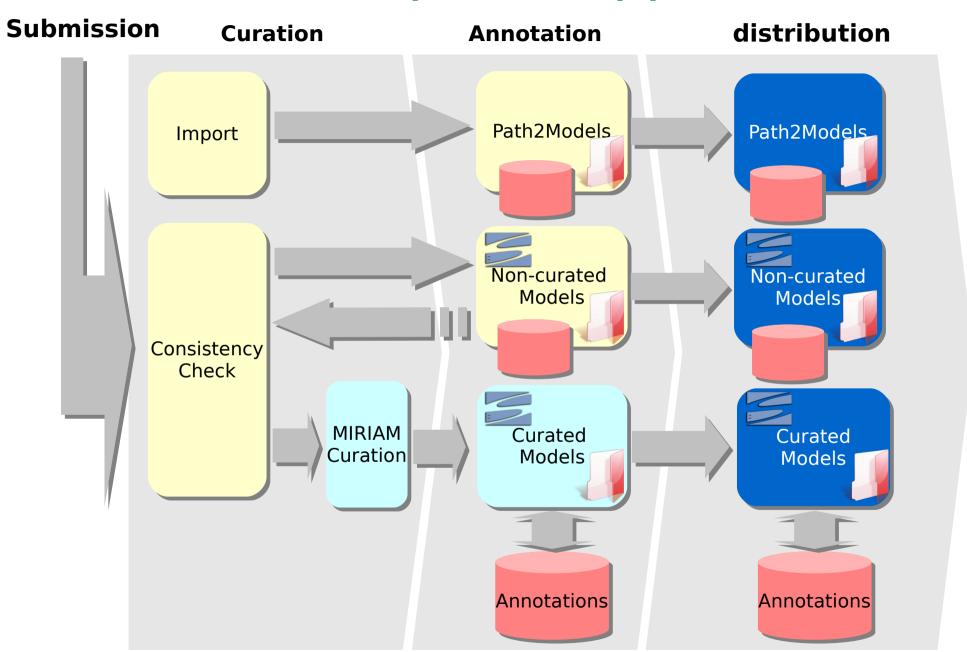


Curated and Non-curated Models

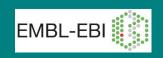
- Curated models Comply with the MIRIAM guidelines
- Non-Curated models valid SBML, not curated or annotated
 - Not MIRIAM compliant:
 cannot reproduce results published in the paper.
 differ in model structure
 non-kinetic models (eg. FBA, stoichiometric maps)
 - MIRIAM compliant:
 models contain kinetics that we cannot curate at present (e.g. reaction-diffusion models)
 models are yet to be curated



Current production pipeline









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BioModels Database

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BioModels Database is a repository of computational models of biological processes. Models described from literature are manually curated and enriched with cross-references. All models are provided in the Public Domain. More information about BioModels Database can be found in the FAO.

Models published in the literature Browse



Manually curated (562 models)



Non curated (696 models) Alternative access



Gene Ontology classification



Gene Ontology tree



Advanced search

Models automatically generated from pathway resources (Path2Models)

Browse



Metabolic (112,898 models) Non-metabolic (27,531 models) Whole genome metabolism (2,641 models) Alternative access







Dedicated search

Taxonomy

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Model of the month

Alpha-synuclein related pathogenic mechanism play a significant role in the development of Parkinson's



disease. In this model, the effects of alphasynuclein oligomerization on cells' homeostasis has been proposed.

Access this model of the month.

Mews



26 February 2015 New CPT:PSP publication

BioModels: Content, Features, Functionality, and

Use published in CPT: Pharmacometrics & Systems Pharmacology

21 November 2014 New NAR publication

BioModels: ten-year anniversary & published in Nucleic Acids Research, Database issue.

11 November 2014 ISBE Business Case

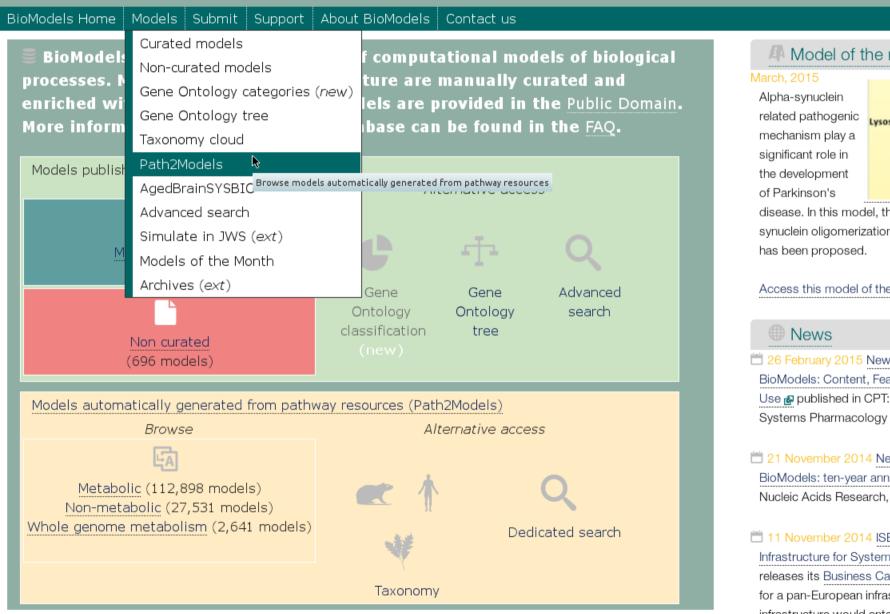
Infrastructure for Systems Biology Europe (ISBE) releases its Business Case, explaining the need for a pan-European infrastructure, what this infrastructure would entail, together with its plans for data stewardship, standardisation and services. BioModels is featured as a success story



BioModels Database

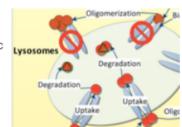
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.ebi.ac.uk/biomodels-main/path2models

Services



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processes. Models

with cross-reference

information about

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SPAROL Endpoint

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Courses

Development with BioModels Database source code ALICHIGUYE GUUESS



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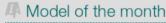




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March, 2015

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with cross-references. All models

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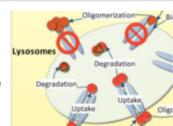
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Submit - Step 1

You can submit here models to be included in BioModels Database.

The following formats are currently accepted:

- SBML Level 3: Version 1 (including any packages)
- SBML Level 2: Version 1, Version 2, Version 3 and Version 4
- SBML Level 1: Version 1 and Version 2

If you wish to submit a model under a different format, please contact us.

If you wish to provide additional files with your submission, please contact us.

The submitted model will not be publicly available from BioModels Database straightaway. If you wish to know more about the <u>submission</u>, <u>curation</u> and <u>annotation</u> processes, please refer to the relevant sections of the <u>Frequently Asked Questions</u>.

To ensure a prompt processing of your model, please follow those simple guidelines:

Check that your model is valid according to the format you chose. If your model is encoded in SBML, you can use the official online validator.

CellML: 1.0 and 1.1

- Enter all relevant information which could help the work of our curators (relation between the model and publication, modifications or clarifications of the model, etc.) either directly into the model file (for example using the *notes* elements if your model is encoded in *SBML*), or in the *Comment* field provided in step 2 of this form.
- If you created the model (or collaborated to its creation) but are not an author of the associated publication, please add your personnal information (first and last name, organisation and email address) in the model, so that your contribution can be acknowledged. If you used SBML, this can be done by adding to the model element a dc:creator annotation, as in this example which you can re-use (skip the blue part if already present).
- Choose a meaningful name for your model. You can follow the pattern AuthorName Year Topic, Method, for example: Levchenko2000 MAPK, noScaffold or Edelstein 1996 EPSP AChEvent.

All models in BioModels Database are available under the terms of the <u>Creative Commons CC0 1.0 (Public Domain Dedication)</u>. Therefore you need to agree to release the encoded model in the **Public Domain** before submitting it to BioModels Database.

Thank you for contributing to BioModels Database!

Please enter the identifier of the scientific publication associated with the model, and then click Continue. If the model has not been published, please select the 'Unpublished' option.

Publication Identifier:

Type of identifier:

• PubMed ID (Search Medline)

• DOI (Resolve a DOI)

• URL

• Unpublished

Continue Reset

Camilana

D----

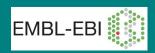
. .

The submitted models will not be incorporated into the BioModels Database straightaway, since they have to undergo a curation phase before. During this curation phase, the models will be first converted to the SBML Level 2 Version 4 format in case they were submitted under a different format, and then tested to verify that they both are consistent and reproduce the results published in the respective reference publication. To actually facilitate this curation phase, prior to submitting a model, please do the following:

- Enter all the relevant information you believe is necessary for the curation (reference publication, modifications or clarifications of the model, etc.) either directly into the model file if allowed (for example using the *notes* elements if your model is under one of the SBML formats), or into the Curation comment text field provided by the form below.
- If you created the model, or collaborated to its creation, and you are not an author of the reference publication, add to the *model* element a *dc:creator* annotation containing your data (first and last name, organisation, email), so that your contribution can be acknowledged. Click here to view an example of a *dc:creator* annotation which you can re-use (skip blue part if already present).
- Choose a meaningful value for the attribute name of the model element. Examples of good model names are NameAuthorYear_Topic_Method, Levchenko2000_MAPK_noScaffold or Edelstein1996 EPSP AChEvent.
- Check the validity of the model (for example by using this online validator if your model is encoded in the SBML format). All the models undergo a primary XML validity check upon submission anyway, and, as mentioned before, a more thorough testing during the curation phase, but an already valid model is of great help nevertheless!
- If the model was not created directly in SBML, or if it requires a specific software to be simulated adequately, please enter in the *Original Model* form a URL pointing to the model in the original repository. Refrain from entering a generic URL to the repository itself.

Please enter your	r personal details and any comment useful for the curation step (underlined fields are requi	equired), and then click <i>Submit</i> .
First name:		
Last name:		
Organisation:		
Email:		
Comment:		
Original model:		
Model file:	Browse No file selected.	
Submit Res	et	
-150		





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Browse - Curated models

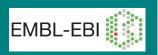
H The following fields are used to describe a model: ...

← 1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23 24 25 26 27 28 29 30 31 32 33 34 35 36 37 38 39 40 41 42 43 44 45 46 47 48 49 50 51 52 53 54 ←

10 | 50 | 100 | All

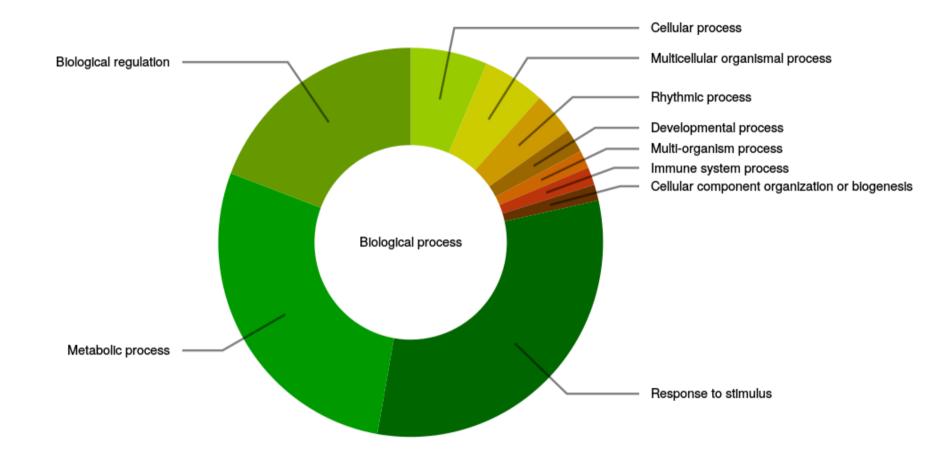
BioModels ID	<u>Name</u> →	Publication ID	<u>Last Modified</u>
BIOMD000000355	Abell2011_CalciumSignaling_WithAdaptation	21844332	2011-09-08T12:03:57+00:00
BIOMD000000354	Abell2011_CalciumSignaling_WithoutAdaptation	21844332	2011-09-08T12:16:51+00:00
BIOMD000000428	Achcar2012 - Glycolysis in bloodstream form T. brucei	22379410	2013-06-10T13:41:07+00:00
BIOMD000000476	Adams2012 - Locke2006 Circadian Rhythm model refined with Input Signal Light Function	22855577	2013-09-11T15:39:47+00:00
BIOMD000000169	Aguda1999_CellCycle	10619492	2012-07-05T14:42:01+00:00
BIOMD000000295	Akman2008_Circadian_Clock_Model1	18277380	2014-05-28T01:56:26+00:00
BIOMD0000000214	Akman2008_Circadian_Clock_Model2	18277380	2014-05-28T01:57:42+00:00
BIOMD0000000220	Albeck2008_extrinsic_apoptosis	18406323	2012-07-05T14:42:40+00:00
BIOMD0000000211	Albert2005_Glycolysis	<u>15955817</u>	2014-05-27T23:52:38+00:00
BIOMD000000289	Alexander2010_Tcell_Regulation_Sys1	20195912	2014-03-11T13:38:50+00:00





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Classification of models published in the literature, based on GO









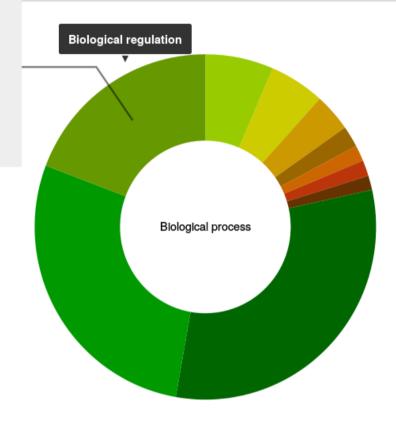
Clas e literature, based on GO

Regulation of biological process (109)

Biological regulation (232)

- · Regulation of membrane potential

- Homeostatic process (27)
 Regulation of localization (21)
 Regulation of molecular function (10)



Selected GO term: Regulation of biological process

Curated Models

BioModels ID	Name	Publication ID	Last Modified
BIOMD000000012	Elowitz2000 - Repressilator	10659856	2013-07-10
BIOMD0000000065	Yildirim2003_Lac_Operon	12719218	2014-02-12
BIOMD0000000067	Fung2005_Metabolic_Oscillator	<u>15875027</u>	2012-05-16
BIOMD0000000079	Goldbeter2006_weightCycling	16595882	2012-07-05
PIOMD000000000	Thomson1090_AdopylatoCycleso	2574002	2014 02 12

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BioModels Database: GO tree

This is a tree view of the curated models based on their Gene Ontology & annotation.

To browse the models, please click the local control is icon on the left of the term to expand the related branch and similarly click on the click on the control collapse an already opened branch. By clicking on a term's name, the list of models annotated with this term (or one of its descendant) will be displayed on the right panel.

- ▼ GO:0008150 biological process (529)
 - ► GO:0022610 biological adhesion (1)
 - ▼ GO:0065007 biological regulation (373)
 - ▼ GO:0050789 regulation of biological process (316)
 - ▼ GO:0048519 negative regulation of biological process (77)
 - ▼ GO:0051129 negative regulation of cellular component organization (5)
 - ▼ GO:0010639 negative regulation of organelle organization (5)
 - ▼ GO:0051784 negative regulation of nuclear division (5)
 - ▼ GO:0045839 negative regulation of mitosis (5)
 - ▼ GO:0045841 negative regulation of mitotic metaphase/anaphase transition (5)

GO:0007094 - mitotic spindle assembly checkpoint (5)

- GO:0048523 negative regulation of cellular process (71)
- GO:0051093 negative regulation of developmental process (1)
- ▶ GO:0045926 negative regulation of growth (1)
- ▶ GO:0002683 negative regulation of immune system process (1)
- ► GO:0009892 negative regulation of metabolic process (54)
- ▶ GO:0043901 negative regulation of multi-organism process (5)
- GO:0051241 negative regulation of multicellular organismal process (1)
- ▶ GO:0048585 negative regulation of response to stimulus (32)
- ▶ GO:0051283 negative regulation of sequestering of calcium ion (7)
- ▶ GO:0023057 negative regulation of signaling (33)
- ► GO:0051051 negative regulation of transport (8)
- ► GO:0048518 positive regulation of biological process (91)
- ▶ GO:0044087 regulation of cellular component biogenesis (2)
- ► GO:0051128 regulation of cellular component organization (12)
- GO:0050794 regulation of cellular process (274)

Model(s) annotated with the GO term $\underline{GO:0007094}$ \mathbf{GO} or one of its descendant(s):

- Chen2004 Cell Cycle Regulation (BIOMD0000000056)
- <u>Ibrahim2008 Mitotic Spindle Assembly Checkpoint Convey variant</u> (BIOMD000000187)
- <u>Ibrahim2008 Mitotic Spindle Assembly Checkpoint Dissociation</u>
 variant (BIOMD000000186)
- <u>Ibrahim2008_Cdc20_Sequestring_Template_Model</u>
 (BIOMD0000000194)
- <u>Ibrahim2008 MCC assembly model KDM</u> (BIOMD000000193)





- Person → Search BioModels Database for model submitter and/or creator(s) names, or model reference publication author(s) names (for example Nicolas L Bruce Shapiro or Shapiro, Edelstein or Novak).
- SBML elements → Search BioModels Database using the content of either "name" or "notes" SBML elements (for example Edelstein or nicotinic). Select the
 you want to find documents which matches the exact phrase; otherwise, all words will be searched as default.
- Annotation (full text) → Search BioModels Database for related information found in the models reference publication or third-party resources, by either publidentifier or text (for example 9256450 or cyclin for publication, GO:0000278 or cell cycle for Gene Ontology, P04551 or cell division for UniProt).
- Annotation (identifier) → Search BioModels Database for annotations, by third-party resource identifiers (for example IPR002394 for InterPro, hsa04080 for 68910 for Reactome).

A part from the *BioModels Identifier* -based search, for every other criteria the search operates on a *contains the entered string basis*, case-insensitive *Person* for *Shapi* or *shapi* will return the same results as searching for *Shapiro* or *shapiro*. In addition, since search strings are treated as words, do n expressions.

Multiple criteria can be combined with either and or or. If and is selected, only those models satisfying all the criteria will be returned. If instead or is smodels satisfying at least one of the criteria will be returned.

		*
		•
BioModels identifier:		
Person:		
SBML elements:		match the exact phrase
Annotation (full text): UniProt Knowledgebase >	Ŷ	
Annotation (full text): Publication cell cycle	住	
Annotation (full text): Gene Ontology	Ġ	1
Annotation (identifier): PubChem-compound		
Annotation (identifier): KEGG Reaction	企	
Annotation (identifier): Enzyme Nomenclature		
Compose by: ● and ○ or		
Search Reset		

Search - Models

Query: Publication cell cycle [resources]

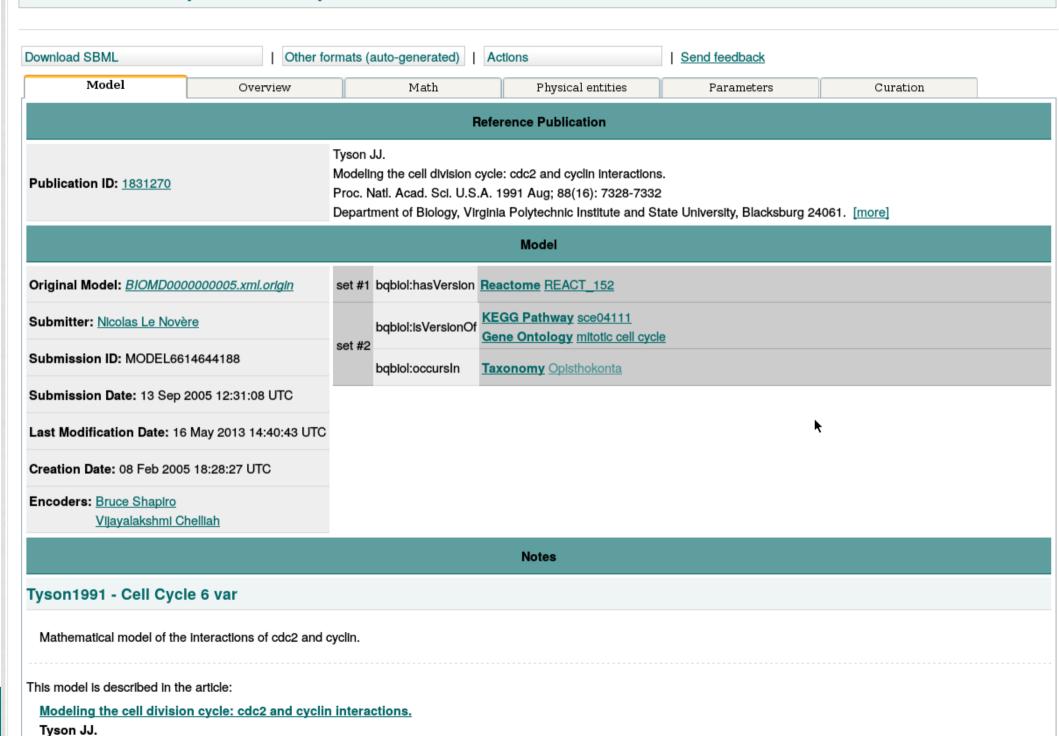
The search returned 44 models.

New Search

■ 33 Curated models returned:

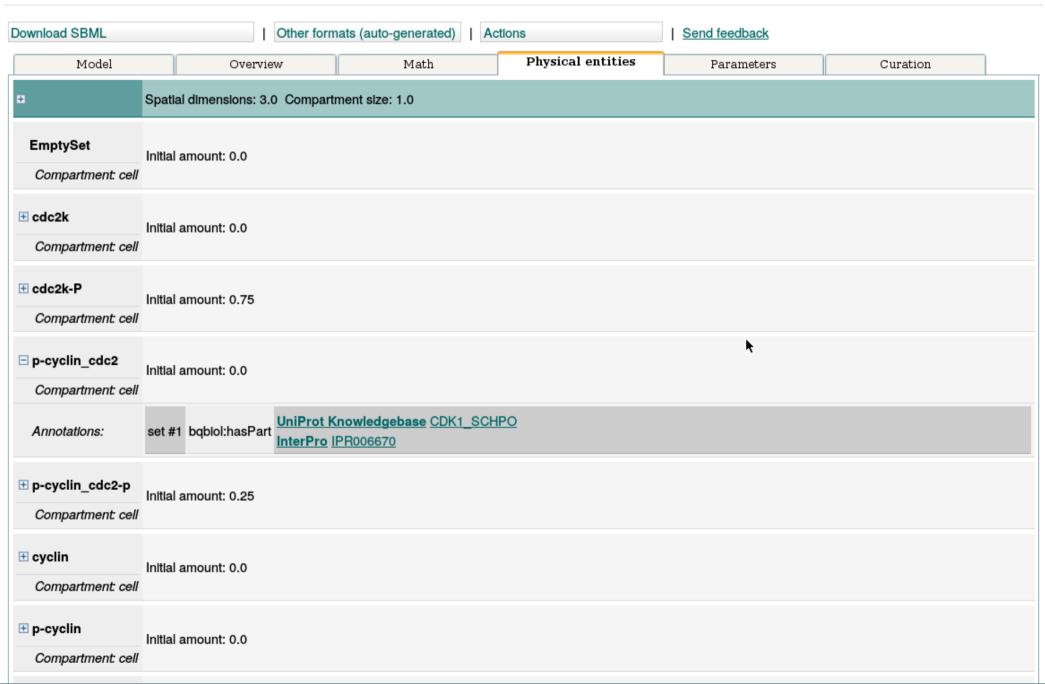
BioModels ID →	<u>Name</u>	Publication ID	<u>Last Modified</u>
BIOMD000000003	Goldbeter1991 - Min Mit Oscil	1833774	2013-05-16T14:38:01+00:00
BIOMD000000004	Goldbeter1991 - Min Mit Oscil, Expl Inact	1833774	2012-12-11T15:30:15+00:00
BIOMD000000005	Tyson1991 - Cell Cycle 6 var	1831270	2013-05-16T14:40:43+00:00
BIOMD000000006	Tyson1991 - Cell Cycle 2 var	1831270	2013-05-16T14:38:56+00:00
BIOMD0000000007	Novak1997 - Cell Cycle	9256450	2014-03-26T14:20:21+00:00
BIOMD000000008	Gardner1998 - Cell Cycle Goldbeter	<u>9826676</u>	2014-07-24T10:59:34+00:00
BIOMD000000018	Morrison1989_FolateCycle	<u>2732237</u>	2012-07-05T14:40:37+00:00
BIOMD000000056	Chen2004 - Cell Cycle Regulation	<u>15169868</u>	2013-06-07T10:59:38+00:00
BIOMD000000109	Haberichter2007_cellcycle	17299420	2012-07-05T14:49:20+00:00
BIOMD000000110	Qu2003_CellCycle	14645053	2012-05-16T10:13:14+00:00
BIOMD000000111	Novak2001_FissionYeast_CellCycle	12779461	2012-07-05T16:47:55+00:00
BIOMD000000150	Morris2002_CellCycle_CDK2Cyclin	11959850	2010-12-01T22:10:11+00:00

BIOMD000000005 - Tyson1991 - Cell Cycle 6 var

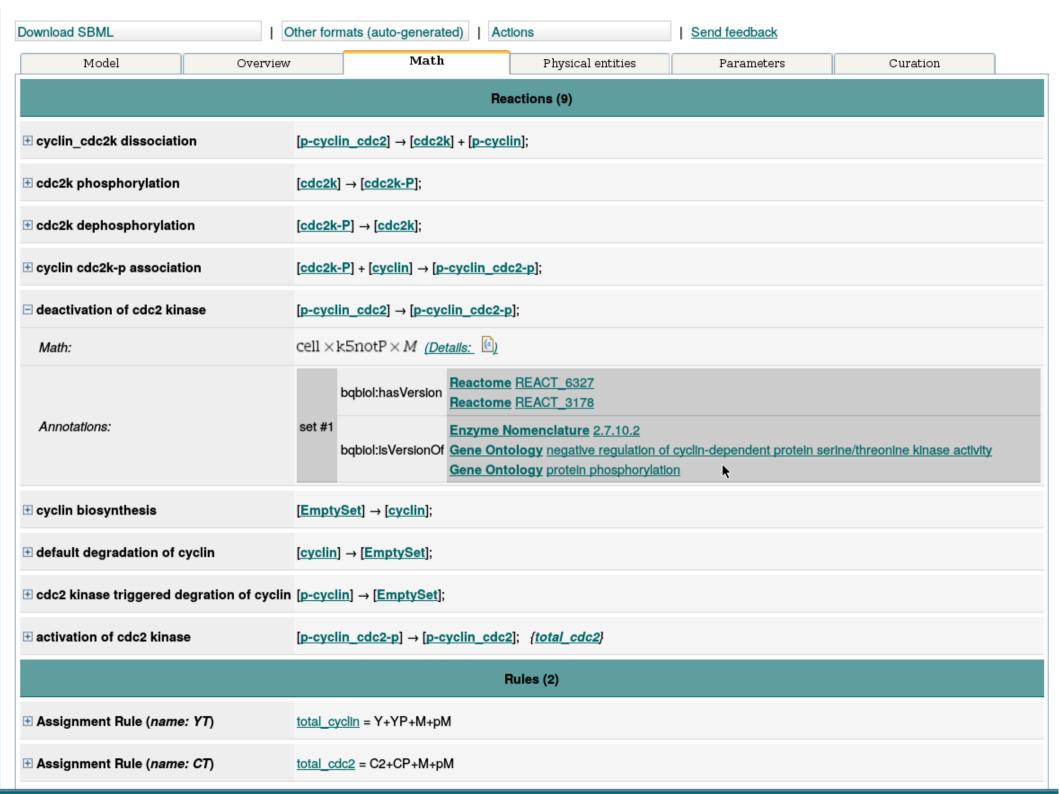


Proc. Natl. Acad. Sci. U.S.A. 1991; 88(16); 7328-32

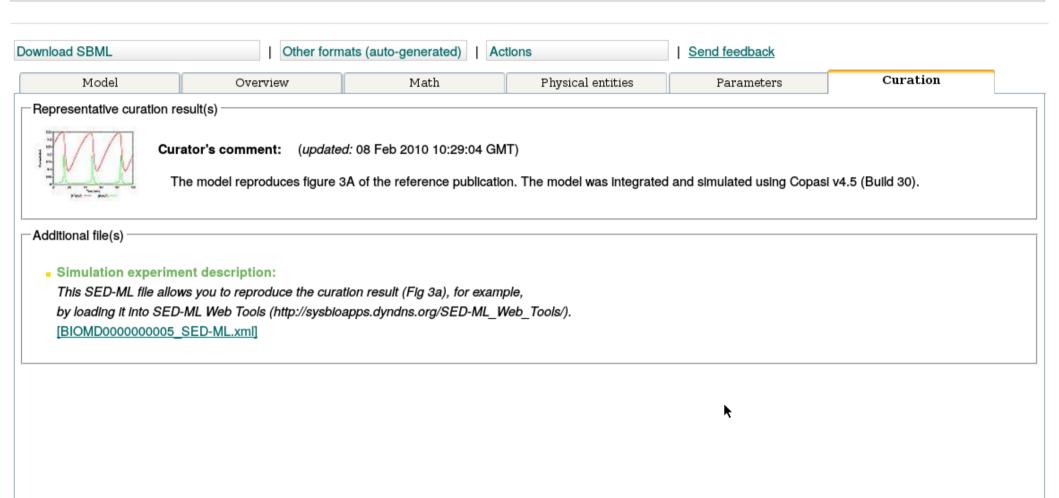
BIOMD000000005 - Tyson1991 - Cell Cycle 6 var



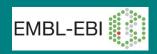




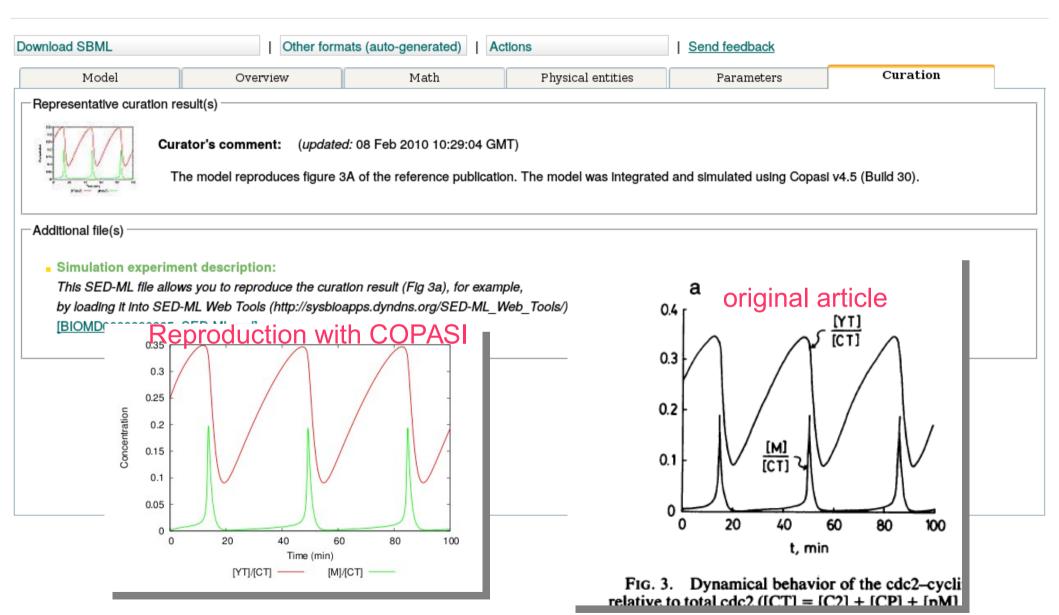
BIOMD000000005 - Tyson1991 - Cell Cycle 6 var







BIOMD000000005 - Tyson1991 - Cell Cycle 6 var



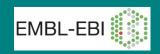


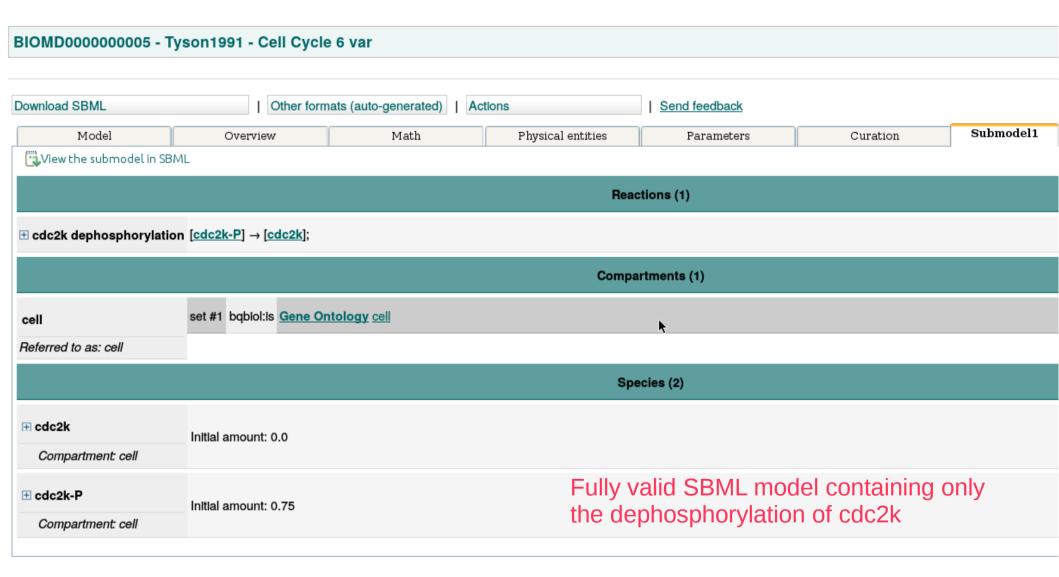
BIOMD00000005 - Tyson1991 - Cell Cycle 6 var								
Download SBML		Other formats (auto-generated)) Actions		Send feedback		
Model Overview Math				Physical entities		Parameters Curation		
ि€Create a submo	del with selected ele	ments					Dese	elect All
				Model				
Publication ID: 18	31270 Submission	Date: 13 Sep 2005 12:31:08 UTC	Las	st Modification Date: 16 May 20	13 14:4	40:43 UTC Creat	tion Date: 08 Feb 2005 18:28:2	27 UTC
			Mathe	ematical expressions				
Reactions								
cyclin_cdc2k dissociation cdc2k phosphorylation cdc2k dephosphorylation cyclin cdc2k-p association								
deactivation of cdc2 kinase cyclin biosynthesis default degradation of cyclin cdc2 kinase triggered degration of cyclin					triggered degration of cyclin	<u>l</u>		
activation of	cdc2 kinase			k				
Rules								
Assignment Rule (variable: total_cyclin) Assignment Rule (variable: total_cdc2)								
Physical entities								
Compartments	Species							
cell	<u>EmptySet</u>		<u> </u>	dc2k			cdc2k-P	
	p-cyclin_cdc2		<u> </u>	-cyclin_cdc2-p			cyclin	
	p-cyclin		<u>to</u>	otal_cyclin			total_cdc2	



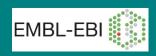
BIOMD00000005 - Tyson1991 - Cell Cycle 6 var									
Download SBML Other formats (auto-generated) Actions Send feedback									
Model	Over	view	Math	Physical entities	Parar	neters	Curatio	n	Submodel1
ि Create a submod	del with selected eleme	ents		-	I			"	Deselect All
				Model					
Publication ID: 1831270 Submission Date: 13 Sep 2005 12:31:08 UTC Last Modification Date: 16 May 2013 14:40:43 UTC Creation Date: 08 Feb 2005 18:28:27 UT					18:28:27 UTC				
			ı	Mathematical expressions					
Reactions									
cyclin_cdc2k c	□ cyclin_cdc2k dissociation □ cdc2k phosphorylation □ cdc2k dephosphorylation □ cyclin cdc2k-p association								
deactivation of cdc2 kinase cyclin biosynthesis			<u>synthesis</u>	default degradation of cyclin cdc2 kinase triggered degration of cyclin			cyclin		
activation of c	activation of cdc2 kinase								
Rules									
Assignment Rule (variable: total_cyclin) Assignment Rule (variable: total_cdc2)									
Physical entities									
□ Compartments □ Species									
cell	EmptySet			cdc2k			cdc2k-P		
p-cyclin_cdc2				p-cyclin_cdc2-p			<u>cyclin</u>		
p-cyclin				total_cyclin			total_cdc2		











BIOMD000000005 - Tyson1991 - Cell Cycle 6 var

Download SBML SBML L2 V1 (auto-generated) SBML L2 V3 (auto-generated) SBML L2 V4 (curated) SBML L3 V1 (auto-generated) SBML L2 V4 (curated with URNs)

Publication ID: 1831270

Automatically generated using libsbml (http://sbml.org/Software/libSBML)

lback

meters

Curation

Reference Publication

Curated version of the model

ate University, Blacksburg 24061. [more]

Model set #1 bqbiol:hasVersion Reactome REACT 152 Original Model: BIOMD000000005.xml.origin KEGG Pathway sce04111 Submitter: Nicolas Le Novère bgbiol:isVersionOf Gene Ontology mitotic cell cycle set #2 Submission ID: MODEL6614644188 babiol:occursIn Taxonomy Opisthokonta

Last Modification Date: 16 May 2013 14:40:43 UTC

Creation Date: 08 Feb 2005 18:28:27 UTC

Submission Date: 13 Sep 2005 12:31:08 UTC

Encoders: Bruce Shapiro

Vijayalakshmi Chelliah

Notes

Tyson1991 - Cell Cycle 6 var

Mathematical model of the interactions of cdc2 and cyclin.

This model is described in the article:

Modeling the cell division cycle: cdc2 and cyclin interactions.

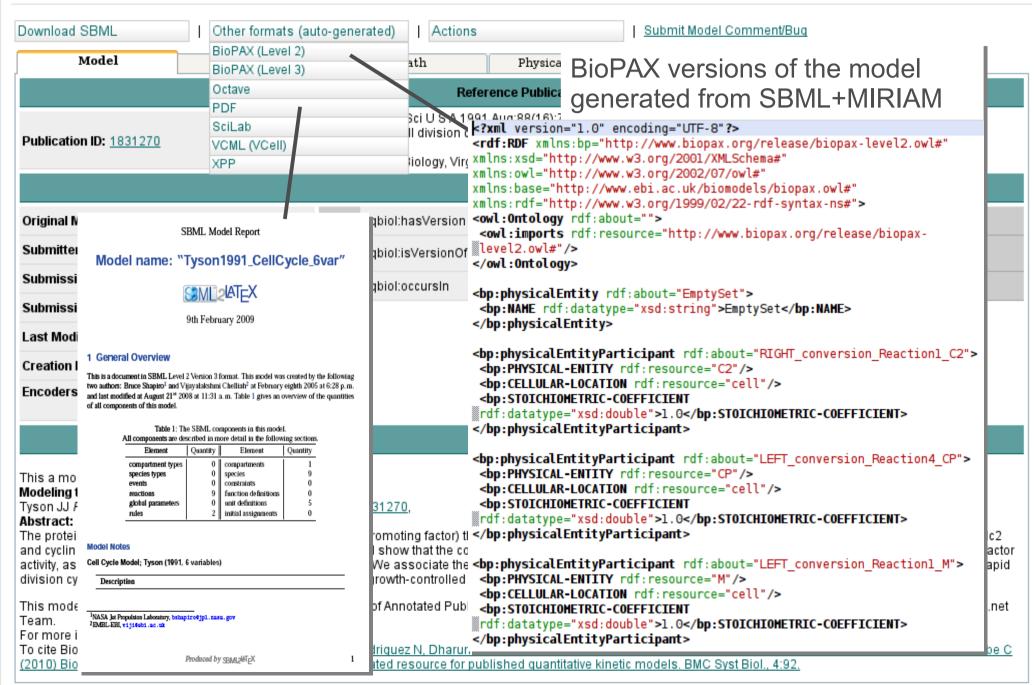
Tyson JJ.

Proc Natl Acad Sci 11 S A 1001 · 88/16) · 7308-30

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BIOMD000000005 - Tyson1991_CellCycle_6var



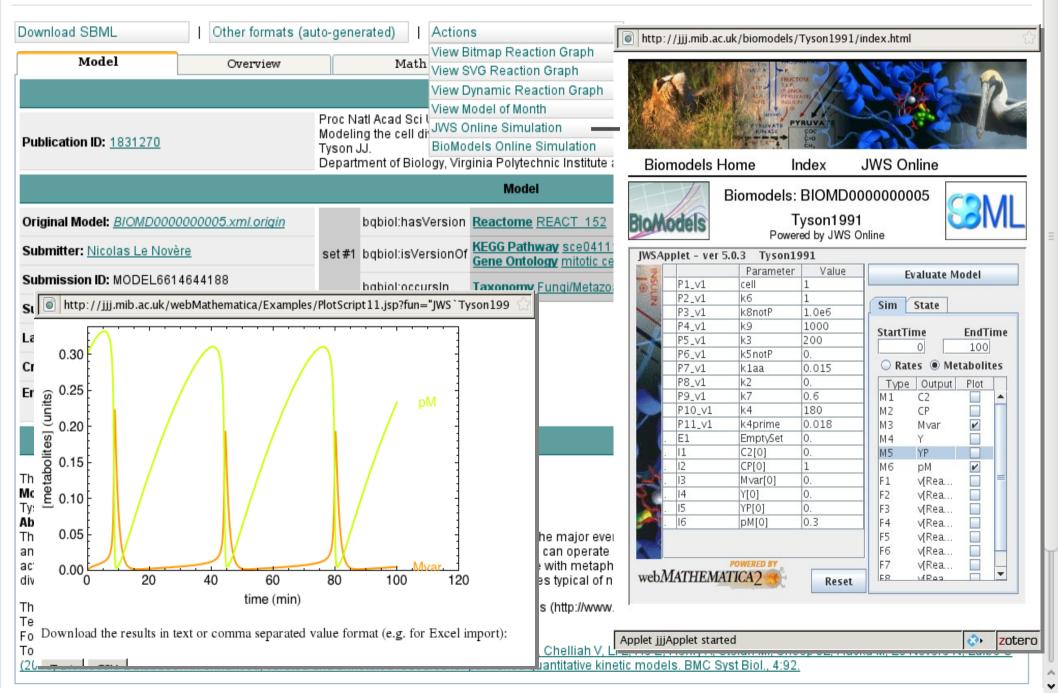


BioModels Home Models Submit Support About BioModels Contact us Search BIOMD000000005 - Tyson1991 CellCycle 6var Download SBML Other formats (auto-generated) Submit Model Comment/Bug Actions View Bitmap Reaction Graph Model Math View SVG Reaction Graph Overview Parameters Curation View Dynamic Reaction Graph View Model of Month Modeling the cell di JWS Online Simulation Proc Natl Acad Sci U Publication ID: 1831270 BioModels Online Simulation Tyson JJ. http://www.ebi.ac.uk/biomodels-main/publ-model-tab.do?cmd=MODEL:SIMU Department of Biology, Virginia Polytechnic Institute and http://www.ebi.ac.uk/biomodels-main/publ-model-tab.do?cmd=MODEL:SIMU:RESULT& Model - Simulation Model The simulation request has been submitted to the queue of our server cluster. Reactome REACT 152 For doing an online simulation, please select the species below. After specifying the simulation time and You could save following links and retrieve your simulation result later. KEGG Pathway sce04111 print step, and then click Submit to submit simulation job to our research cluster. Link of simulation result: Gene Ontology mitotic cell cycl Click Cancel to close the window http://www.ebi.ac.uk/biomodels-main/publ-model.do?cmd=SIMU:RETRIEVE& Faxonomy Fungi/Metazoa grou simuid=SIMU1234453623040 Cancel Species ✓ cdc2k-P ☐ EmptySet cdc2k p-cyclin cdc2-p cyclin p-cyclin 0.7 total cdc2 0.6 Notes Simulation Time (use scientific notation e.g. 1e7 for 10000000): 100 Print step: 1000 0.5 Submit 0.4 at controls the major events of trol system can operate in thre 0.2 steady state with metaphase a ivision cycles typical of nonem 0.1 shed Models (http://www.ebi.ac 100 H, Endler L, Chelliah V, Li L, H Close ublished quantitative kinetic m Done **⊘**⊢ zotero http://www.ebi.ac.uk/biomodels/models-main/ode_simu/SIMU1234453623. zotero

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BIOMD000000005 - Tyson1991_CellCycle_6var





From pathways to models ... Path2Models





PathwayInteractionDatabase



Logical models of individual signalling pathways





Chemical kinetics models of individual metabolic pathways







Flux Balance Analysis of whole genome reconstructions





Path2Models

Path2Models is a branch of BioModels Database dedicated to hosting models automatically generated from pathway resources, such as <u>KEGG</u>, <u>BioCarta</u>, <u>MetaCyc</u>, <u>PID</u> and <u>SABIO-RK</u>.

Browse models

Models from this branch are classified in 3 distinct categories:

- · metabolic models
- non-metabolic models
- · whole genome metabolism models

One can also browse those models by organism:

list of all organisms

Search models

The following search will only look for models coming from the path2models project:

N

Help about the search

- The keywords AND or OR (in upper cases) are available to refine the search. By default, if more than one word is present in a query, OR will be used to combine them.
- Double quotes (") can be used to force the search engine to match a whole expression containing several words.

Search

• The colon character (:) must be escaped in the queries; one can use a backslash for this purpose (1:).

Download all models

• Archives of all models (from the latest release)

Additional information

Those automatically generated models are only partially parameterized. In the case of KEGG signaling pathways for which no mechanistic details are provided, the models (with qual constructs) contain only topological relationships together with interaction signs. No logical rules specify the effects of (combined) interactions, and these models should be seen as confident to be further parameterized before use in simulation. This can be done either by considering default, yet biologically meaningful, logical functions

Path2Models:

Here are all the whole genome metabolism models available:

- [Eubacterium] cylindroides T2-87
- Acaryochloris marina (strain MBIC 11017)
- Accumulibacter phosphatis (strain UW-1)
- Acetobacter pasteurianus (strain NBRC 3283 / LMG 1513 / CCTM 1153)
- Acetobacter pasteurlanus IFO 3283-01-42C
- Acetobacter pasteurianus IFO 3283-03
- Acetobacter pasteurlanus IFO 3283-07
- Acetobacter pasteurianus IFO 3283-12
- Acetobacter pasteurianus IFO 3283-22
- Acetobacter pasteurlanus IFO 3283-26
- Acetobacter pasteurianus IFO 3283-32
- Acetobacterium woodii (strain ATCC 29683 / DSM 1030 / JCM 2381 / KCTC 1655)
- Acetohalobium arabaticum (strain ATCC 49924 / DSM 5501 / Z-7288)
- Acholeplasma laidlawii (strain PG-8A)
- Achromobacter xylosoxidans (strain A8)
- Acidaminococcus fermentans (strain ATCC 25085 / DSM 20731 / VR4)
- Acidaminococcus intestini (strain RyC-MR95)
- Acidianus hospitalis (strain W1)
- Acidilobus saccharovorans (strain DSM 16705 / VKM B-2471 / 345-15)
- Acidimicrobium ferrooxidans (strain DSM 10331 / JCM 15462 / NBRC 103882 / ICP)
- Acidiphilium cryptum (strain JF-5)
- Acidiphilium multivorum (strain DSM 11245 / JCM 8867 / AIU301)
- Acidithiobacillus caldus (strain SM-1)
- Acidithiobacillus ferrivorans SS3
- Acidithiobacillus ferrooxidans (strain ATCC 23270 / DSM 14882 / NCIB 8455)
- Acidithiobacillus ferrooxidans (strain ATCC 53993)
- Acidobacterium capsulatum (strain ATCC 51196 / DSM 11244 / JCM 7670)
- Acidothermus cellulolyticus (strain ATCC 43068 / 11B)
- Acidovorax avenae (strain ATCC 19860 / DSM 7227 / JCM 20985 / NCPPB 1011)
- Acidovorax citrulli (strain AAC00-1)
- Acidovorax ebreus (strain TPSY)
- Acidovorax sp. (strain JS42)

•

Whole Genome Metabolism - Homo sapiens

Homo sapiens

Download SBML

SBML L2 V4

SBML L2 V4 (with Identifiers.org URLs)

Send feedback

Model information

Identifier: BMID000000140905

Format: SBML L2 V4

occursin

Project:

path2models

Submission: 19 May 2012 15:48:00 UTC

Categories: genome-scale

Last modified: 04 Oct 2013 04:05:27 UTC

Taxonomy

Published: 19 May 2012 23:49:21 UTC

Annotations

	occursiii	Tionio sapiens	Taxoriomy
	IsDerivedFrom	Homo saplens (human)	KEGG Genome
	IsDescribedBy	Initial sequencing and analysis of the human genome.	PubMed
	isDescribedBy	The sequence of the human genome.	PubMed
	isDescribedBy	Finishing the euchromatic sequence of the human genome.	PubMed
	isDescribedBy	The DNA sequence and biological annotation of human chromosome 1.	PubMed
	isDescribedBy	Generation and annotation of the DNA sequences of human chromosomes 2 and 4.	PubMed
	isDescribedBy	The DNA sequence, annotation and analysis of human chromosome 3.	PubMed
	isDescribedBy	The DNA sequence and comparative analysis of human chromosome 5.	PubMed
	isDescribedBy	The DNA sequence and analysis of human chromosome 6.	PubMed
	isDescribedBy	Human chromosome 7: DNA sequence and biology.	PubMed
	isDescribedBy	The DNA sequence of human chromosome 7.	PubMed
	isDescribedBy	DNA sequence and analysis of human chromosome 8.	PubMed
	isDescribedBy	DNA sequence and analysis of human chromosome 9.	PubMed
	isDescribedBy	The DNA sequence and comparative analysis of human chromosome 10.	PubMed
	isDescribedBy	Human chromosome 11 DNA sequence and analysis including novel gene identification.	PubMed
	isDescribedBy	The finished DNA sequence of human chromosome 12.	PubMed
	isDescribedBy	The DNA sequence and analysis of human chromosome 13.	PubMed
	isDescribedBy	The DNA sequence and analysis of human chromosome 14.	PubMed
	isDescribedBy	Analysis of the DNA sequence and duplication history of human chromosome 15.	PubMed
ļ	isDescribedBy	The sequence and analysis of duplication-rich human chromosome 16.	PubMed

BioModels Web Services

With BioModels Web Services, users can programmatically access up-to-date information from BioModels Database without installing a local copy of the database. They provide a wide range of features for searching and retrieving models. Furthermore, some features can help users to extract interesting parts from a large model and assemble them into a fully valid submodel. For any comments or new feature enquiries, please feel free to contact us.

Available features

- Available features
- WSDL

The list of available features's page describes all the available services in a nice human readable way, included a detailed description of all methods. The WSDL (Web Services Description Language) defines the provided services in an XML format file. This enables third-party software to automatically generate clients for accessing the services.

Java library

The Java library provides a very convenient way to use the web services. It gives access to improved methods (for example giving access to 'SimpleModel' objects rather than raw XML) in order to make the use of the web services easier.

k

Documentation

The library documentation gives information for developers wishing to use the API:

Java library documentation (Javadoc)

Get the library via Maven

The library is available from the EBI Maven repository.

You just need to add the following to your pom.xml to get it and all its dependencies:

Index of ftp://ftp.ebi.ac.uk/pub/databases/biomodels/weekly_archives/2012/

♠ Up to higher level directory

Name	Size	Last Modified		
p BioModels-Database-weekly-2012-01-02-sbmls.tar.gz	11734 KB	22/03/12 00:00:00		
p BioModels-Database-weekly-2012-01-09-sbmls.tar.gz	11735 KB	22/03/12 00:00:00		
BioModels-Database-weekly-2012-01-16-sbmls.tar.gz	11734 KB	22/03/12 00:00:00		

Index of ftp://ftp.ebi.ac.uk/pub/databases/biomodels/releases/

♣ Up to higher level directory

Name	Size	Last Modified	
2005-04-11		08/02/12	00:00:00
2005-06-01		08/02/12	00:00:00
2005-07-28		08/02/12	00:00:00
2006-01-31		08/02/12	00:00:00

Index of ftp://ftp.ebi.ac.uk/pub/databases/biomodels/releases/latest/

♠ Up to higher level directory

Name	Size	Last Modified		
→ BioModels_Database-r23_p2m-metabolic.tar.bz2	1966032 KB	10/08/12 15:19:00		
── BioModels_Database-r23_p2m-non_metabolic.tar.bz2	78726 KB	10/08/12 15:19:00		
😿 BioModels_Database-r23_p2m-whole_genome_metabolism.tar.bz2	509109 KB	10/08/12 15:20:00		
₩ BioModels_Database-r23_pub-all_files.tar.bz2	601968 KB	10/08/12 21:03:00		
₩ BioModels_Database-r23_pub-sbml_files.tar.bz2	10992 KB	11/08/12 13:10:00		



EMBL-EBI

RDF Platform

RDF Platform

Services

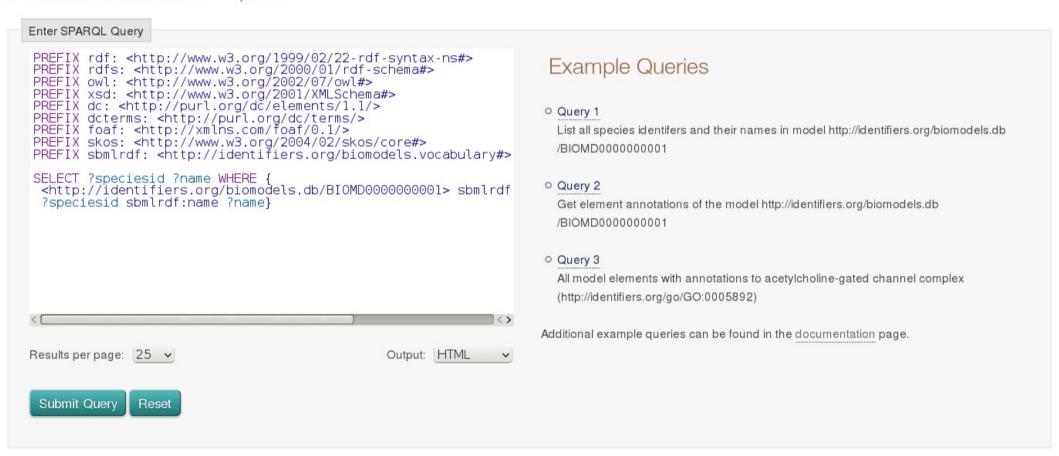
Documentation

FAQ About



Services > BioModels > BioModels SPAROL Endpoint

BioModels SPARQL Endpoint





EMBL-EBI

Developers

Mélanie Courtot, Alexander Broicher, Finja Büchel, Marco Donizelli, Marine Dumousseau, Gael Jalowicki, Mihai Glont, Ron Henkel, Arnaud Henri, Sarah Keating, Christian Knuepfer, Camille Laibe, Chen Li, Lu Li, Florian Mittag, Kedar Nath Natarajan, Jean-Baptiste Pettit, Nicolas Rodriguez, Karim Tazibt, Martijn van Iersel, Dagmar Waltemath, Sarala Wimalaratne, Yangyang Zhao, Anna Zhukova

Curators

Ishan Ajmera, Vijayalakshmi Chelliah, Harish Dharuri, Lukas Endler, Enuo He, Nick Juty, Vincent Knight-Schrijver, Audald Lloret, Michael Schubert, Melanie Stefan

Henning Hermjakob, Janet Thornton

External contributions

Collaborators

Upi Bhalla, Andreas Dräger, Jürgen Eils, Mike Hucka, Hiroaki Kitano, Ion Moraru, Julio Saez-Rodriguez, Herbert Sauro, Maria Schilstra, Falk Schreiber, Jacky Snoep, Neil Swainston

- Scientific Advisory Board
 Upi Bhalla, Carole Goble, Thomas
 Lemberger, Pedro Mendes, Ion
 Moraru, Wolfgang Müller, Philippe
 Sanseau, Herbert Sauro, Jacky Snoep
- All the computational systems Biology community, in particular: Rainer Machne, Bruce Shapiro, Kieran Smallbone

Current member coordinator





Generous funders















