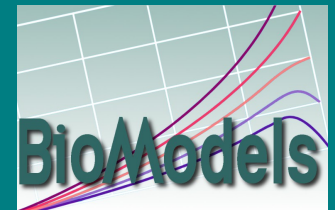


BioModels Database

a database of quantitative, kinetic models



CCB 2010
Hinxton, UK

Lukas Endler
<lukas@ebi.ac.uk>



EBI is an Outstation of the European Molecular Biology Laboratory.

BioModels Database

- contains only models from the peer reviewed literature
- unique perennial identifiers for models
 - can be referenced, eg. in publications
- models freely accessible and reusable
- models are manually curated and checked to ensure reliability
 - curated and non curated branch
- models and model elements cross-linked to and annotated with controlled vocabularies and databases
 - allows for complex queries and detailed searching
 - adds information and eases identification of model elements

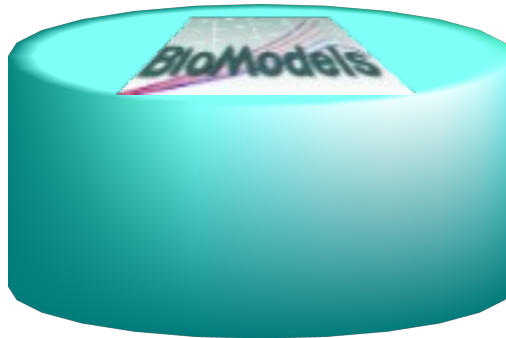
* MIRIAM: Minimal Information Required In the Annotation of Biochemical Models
Nicolas Le Novère et al., *Nature Biotechnology*, **23**(12), 2005



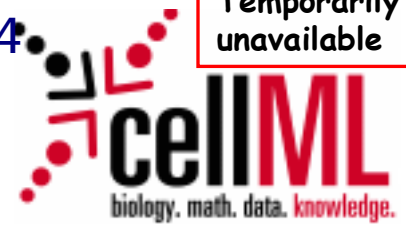
Level 1 Version 1
Level 1 Version 2
Level 2 Version 1
Level 2 Version 3
Level 2 Version 4



Version 1.0
Version 1.1



Level 2 Version 1
Level 2 Version 3
Level 2 Version 4



Temporarily
unavailable

Version 1.1



XPP-Aut



VCell

BioPAX



Model Submission

Where do models come from?

- submitted by curators
 - from other repositories (JWS online, DOQCS, VCell and CellML repositories, ...)
 - reimplemented from literature
 - from journals webpages
- from authors before publication
 - some journals advocate submission to BioModels DB:
 - Molecular Systems Biology
 - PLoS journals
 - BioMedCentral journals
- various people that deem the model to be of interest

Methodology article

Open Access

Towards a genome-scale kinetic model of cellular metabolism

Kieran Smallbone , Evangelos Simeonidis , Neil Swainston  and Pedro Mendes 

BMC Systems Biology 2010, **4**:6 doi:10.1186/1752-0509-4-6

Published: 28 January 2010

Abstract (provisional)

Background

Advances in bioinformatic techniques and analyses have led to the availability of genome-scale metabolic reconstructions. The size and complexity of such networks often means that their potential behaviour can only be analysed with constraint-based methods. Whilst requiring minimal experimental data, such methods are unable to give insight into cellular substrate concentrations. Instead, the long-term goal of systems biology is to use kinetic modelling to characterize fully the mechanics of each enzymatic reaction, and to combine such knowledge to predict system behaviour.

Results

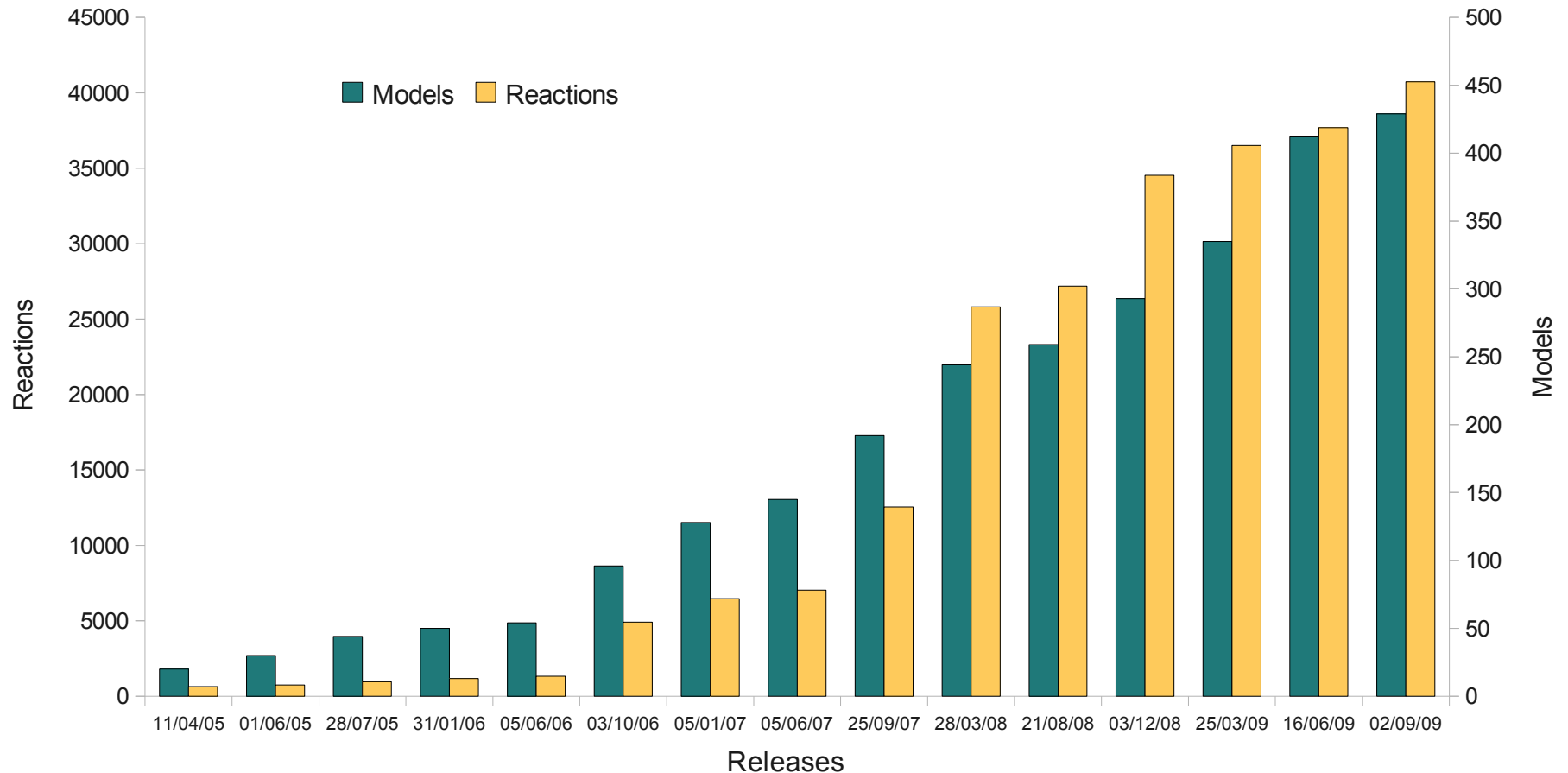
We describe a method for building a parameterized genome-scale kinetic model of a metabolic network. Simplified linlog kinetics are used and the parameters are extracted from a kinetic model repository. We demonstrate our methodology by applying it to yeast metabolism. The resultant model has 956 metabolic reactions involving 820 metabolites, and, whilst approximative, has considerably broader remit than any existing models of its type. Control analysis is used to identify key steps within the system.

Conclusions

Our modelling framework may be considered a stepping-stone toward the long-term goal of a fully-parameterized model of yeast metabolism. The model (see additional file 1) is available in SBML format from the BioModels database (BioModels ID:

MODEL1001200000) and at <http://www.mcisb.org/resources/genomescale/>.

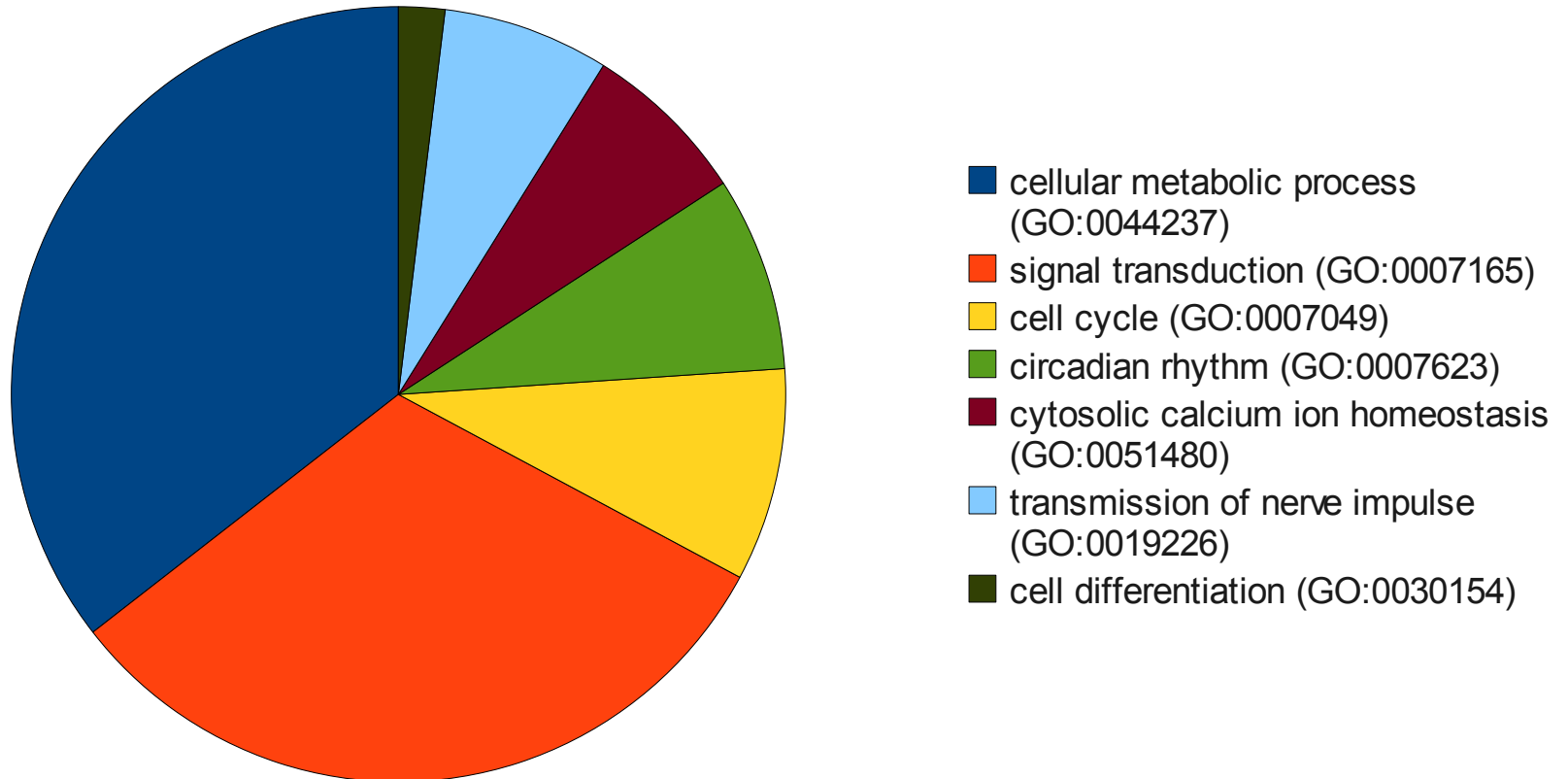
Models in BioModels DB



6

curated branch : 241
non-cura. branch: 213

Models in BioModels DB



Curated and Noncurated Branch

Curated models

- models reproduce results, fully annotated, MIRIAM compliant

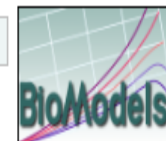
Non-Curated models

- valid SBML, not curated or annotated by the curators.
 - not MIRIAM compliant
 - can not reproduce results published in the paper.
 - non kinetic models (eg. FBA, stoichiometric maps).
 - MIRIAM compliant
 - models contain kinetics that we cannot curate up to now.
 - back lag in curation, the curators just did not have the time → these models will be moved into the curated branch as soon as possible.

http://www.ebi.ac.uk/biomodels

BioModels Database - A Database of Annotated Published Models

BioModels Database is a data resource that allows biologists to store, search and retrieve published mathematical models of biological interests. Models present in BioModels Database are annotated and linked to relevant data resources, such as publications, databases of compounds and pathways, controlled vocabularies, etc.



[Advanced search](#)

Browse models

- [Curated models \(241\)](#)
- [Browse models using GO](#)
- [Non-curated models \(212\)](#)

Simulate in JWS Online

Submit a model

Mirror at California Institute of Technology <http://biomodels.caltech.edu>

BioModels AT SourceForge <http://sourceforge.net/projects/biomodels/>

Web Services <http://www.ebi.ac.uk/biomodels-main/webservices>

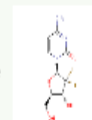
Download archived models <http://www.ebi.ac.uk/biomodels/models-main/tars/>

Model of the month

January, 2010

The pharmacodynamic model deducing the potency and time course of the anticancer agent in shrinking the primary tumours can have potential role to decide dosage and treatment duration.

[Read more...](#)



News

26th January 2010 **Sixteenth release**

[Download All Models Under SBML Format](#)

2nd September 2009 **Fifteenth release**

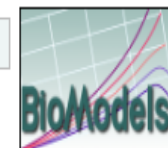
[Download All Models Under SBML Format](#)

16th June 2009 **Fourteenth release**

[Download All Models Under SBML Format](#)

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[Advanced search](#)

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BioModels AT SourceForge <http://sourceforge.net/projects/biomodels/>

Web Services <http://www.ebi.ac.uk/biomodels-main/webservices>

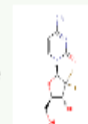
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Browse - Curated models



This is a tree view of the models in BioModels Database based on [Gene Ontology](#). To browse the models, please click to expand the branch, or click to collapse the branch. By double clicking the Gene Ontology term, the detail of the term will be displayed in a new window. By double clicking the BioModels Model ID, this page will be forwarded to the detail of selected model.

GO:0008150 - biological_process (230)

GO:0005575 - cellular_component (200)

GO:0003674 - molecular_function (154)

BioModels ID: *Unspecified*

Name: *N/A*

Publication ID: *N/A*

Last Modified: *N/A*

The relationships between terms are represented by different icons.

- BioModels qualifiers:
 - bqbiol:is
 - bqbiol:isVersionOf
 - bqbiol:hasPart
- Gene Ontology relationships:
 - is a
 - part of
 - develops from
 - other

Browse - Curated models



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- GO:0008150 - biological_process (230)
 - GO:0009987 - cellular process (213)
 - GO:0051641 - cellular localization (45)
 - GO:0050794 - regulation of cellular process (141)
 - GO:0007049 - cell cycle (23)
 - GO:0051726 - regulation of cell cycle (19)
 - GO:0000278 - mitotic cell cycle (21)
 - GO:0051329 - interphase of mitotic cell cycle (5)
 - GO:0000087 - M phase of mitotic cell cycle (2)
 - GO:0007346 - regulation of mitotic cell cycle (10)
 - GO:0051439 - regulation of ubiquitin-protein ligase activity during mitotic cell cycle (4)
 - GO:0045931 - positive regulation of mitotic cell cycle (1)
 - GO:0007052 - mitotic spindle organization (1)
 - BIOMD0000000003 - Goldbeter1991_MinMitOscil
 - BIOMD0000000004 - Goldbeter1991_MinMitOscil_Explnact
 - BIOMD0000000005 - Tyson1991_CellCycle_6var**
 - BIOMD0000000006 - Tyson1991_CellCycle_2var
 - BIOMD0000000007 - Novak1997_CellCycle
 - BIOMD0000000008 - Gardner1998_CellCycle_Goldbeter
 - BIOMD0000000056 - Chen2004_CellCycle
 - BIOMD0000000069 - Fuss2006_MitoticActivation
 - BIOMD0000000107 - Novak1993_M_phase_control
 - BIOMD0000000110 - Qu2003_CellCycle
 - BIOMD0000000111 - Novak2001_FissionYeast_CellCycle
 - BIOMD0000000144 - Calzone2007_CellCycle
 - BIOMD0000000150 - Morris2002_CellCycle_CDK2Cyclin
 - BIOMD0000000168 - Obeyesekere1999_CellCycle
 - BIOMD0000000181 - Sriram2007_CellCycle
 - BIOMD0000000207 - Romond1999_CellCycle
 - BIOMD0000000208 - Deinek02003_CellCycle
 - GO:0022402 - cell cycle process (9)
 - GO:0045786 - negative regulation of cell cycle (14)
 - GO:0045787 - positive regulation of cell cycle (11)
 - BIOMD0000000196 - Srividhya2006_CellCycle
 - GO:0022402 - cell cycle process (9)
 - GO:0016043 - cellular component organization (67)
 - GO:0007154 - cell communication (98)

BioModels ID: [BIOMD0000000005](#)

Name: Tyson1991_CellCycle_6var

Publication ID: [1831270](#)

Last Modified: 2009-08-10T14:09:39+00:00

SBML L2 V4

Complex Searches

Search - Models

 [Text Search](#)

You can search BioModels Database for models using one or more of the following criteria:

- **BioModels ID** → Search BioModels Database for *exact* BioModels identifiers (for example *BIOMD0000000001* or *BIOMD0000000022*).
- **Person** → Search BioModels Database for model submitter and/or creator(s) names, or model reference publication author(s) names (for example *Nicolas Le Novère*, *Nicolas*, *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 checkbox behind, if you want to find documents which matches the exact phrase; otherwise, all words will be searched as default.
- **Resource** → Search BioModels Database for related information found in the models reference publication or third-party resources, by either publication/resource identifier or text (for example *9256450* or *cyclin* for publication, *GO:0000278* or *cell cycle* for [Gene Ontology](#), *P04551* or *cell division* for [UniProt](#)).
- **Resource ID** → Search BioModels Database for annotations, by third-party resource identifiers (for example *IPR002394* for [InterPro](#), *hsa04080* for [KEGG Pathway](#), *68910* for [Reactome](#)).

A part from the *BioModels ID*-based search, for every other criteria the search operates on a *contains the entered string basis*, case-insensitive. That is, searching *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 not enter regular 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 selected, all the models satisfying at least one of the criteria will be returned.

BioModels ID:

Person:

SBML Elements: ☐ match the exact phrase

Resource: 

Resource: 

Resource: 

Resource ID: 

Resource ID: 

Resource ID: 

Compose by: ☒ and ☐ or

BIOMD0000000005 - Tyson1991_CellCycle_6var



SBML formats
 |
 Other formats
 |
 Actions
 |
 [Submit Model Comment/Bug](#)

Model	Overview	Math	Physical entities	Parameters	Curation
-------	----------	------	-------------------	------------	----------

Reference Publication

Publication ID: [1831270](#)
 Proc Natl Acad Sci U S A 1991 Aug;88(16):7328-32.
 Modeling the cell division cycle: cdc2 and cyclin interactions.
 Tyson JJ.
 Department of Biology, Virginia Polytechnic Institute and State University, Blacksburg 24061.
 [\[more\]](#)

Model

Original Model: BIOMD0000000005.xml.orig.in	set #1	bqbiol:hasVersion	Reactome REACT_152
Submitter: Nicolas Le Novère		bqbiol:isVersionOf	KEGG Pathway sce04111 Gene Ontology mitotic cell cycle
Submission ID: MODEL6614644188		bqmodel:is	Taxonomy Funqi/Metazoa group
Submission Date: 13 Sep 2005 12:31:08 UTC			
Last Modification Date: 10 Aug 2009 14:09:39 UTC			
Creation Date: 08 Feb 2005 18:28:27 UTC			
Encoders: Bruce Shapiro Vijayalakshmi Chelliah			

Notes

This a model from the article:
Modeling the cell division cycle: cdc2 and cyclin interactions.
 Tyson JJ *Proc. Natl. Acad. Sci. U.S.A.*1991; 88(16); 7328-32 [1831270](#).
Abstract:
 The proteins cdc2 and cyclin form a heterodimer (maturation promoting factor) that controls the major events of the cell cycle. A mathematical model for the interactions of cdc2 and cyclin is constructed. Simulation and analysis of the model show that the control system can operate in three modes: as a steady state with high maturation promoting factor activity, as a spontaneous oscillator, or as an excitable switch. We associate the steady state with metaphase arrest in unfertilized eggs, the spontaneous oscillations with rapid division cycles in early embryos, and the excitable switch with growth-controlled division cycles typical of nonembryonic cells.
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 For more information see the [terms of use](#).
 To cite BioModels Database, please use [Le Novère N., Bornstein B., Broicher A., Courtot M., Donizelli M., Dharuri H., Li L., Sauro H., Schilstra M., Shapiro B., Snoep J.L., Hucka M. \(2006\) BioModels Database: A Free, Centralized Database of Curated, Published, Quantitative Kinetic Models of Biochemical and Cellular Systems Nucleic Acids Res. 34: D689-D691.](#)

SBML formats | Other formats | Actions | [Submit Model Comment/Bug](#)

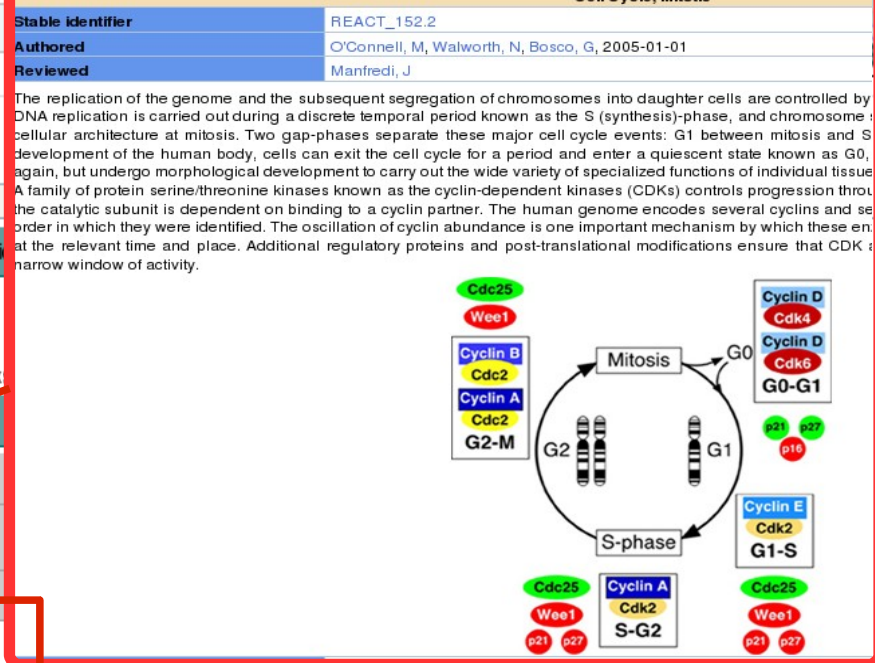
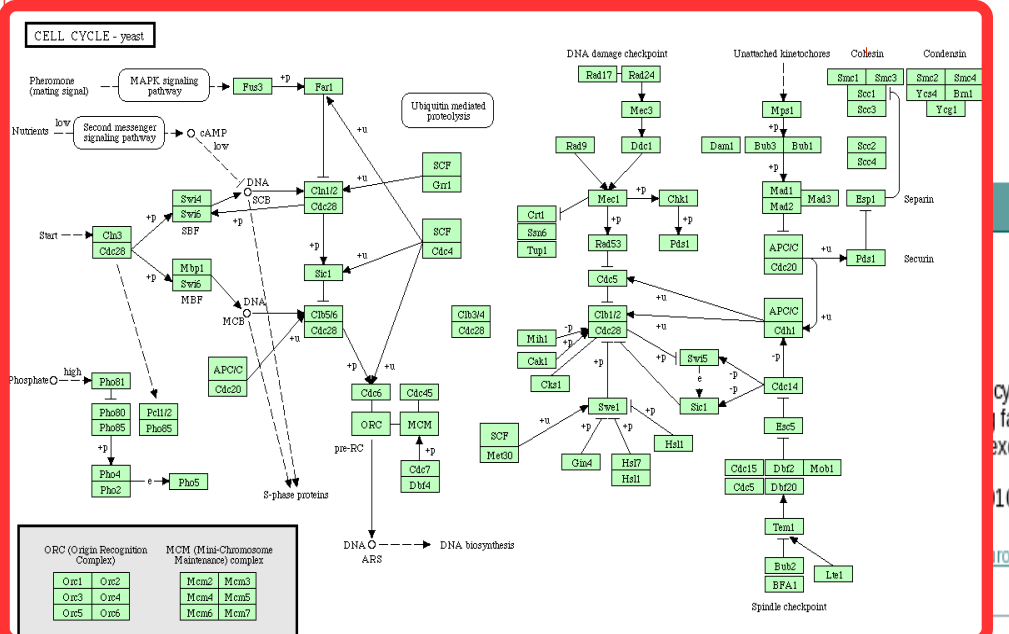
Model | Overview | Math | Physical entities | Parameters

Reference Publications

Publication ID: [1831270](#)
Proc Natl Acad Sci U S A 1991 Aug;88(16):7328-32.
Modeling the cell division cycle: cdc2 and cyclin interactions.
Tyson JJ.
Department of Biology, Virginia Polytechnic Institute and State University, Blacksburg, VA 24061-0400, USA

Original Model: [BIOMD0000000005.xml.orig.in](#)
Submitter: [Nicolas Le Novère](#)
Submission ID: MODEL661464418
Submission Date: 13 Sep 2005 12:31:08 UTC

bqbiol:hasVersion [Reactome REACT_152](#)
set#1 bqbiol:isVersionOf [KEGG Pathway sce04111](#)
[Gene Ontology mitotic cell cycle](#)
bqmodel:is [Taxonomy Fungi/Metazoa group](#)



Lineage	Taxonomy identifier	33154
	Scientific name	Fungi/Metazoa group
	Common name	-
• Eukaryota	Other NCBI synonyms	Opisthokonta opisthokonts
	Rank	no rank
	Number of UniProtKB/Swiss-Prot entries	114913
	Number of UniProtKB/TrEMBL entries	1459130

Taxonomy navigation	
Up taxonomy tree	Down taxonomy tree
Eukaryota	• Choanoflagellida • Fungi • Fungi/Metazoa incertae sedis • Metazoa

BIOMD0000000005 - Tyson1991_CellCycle_6var



SBML formats | Other formats | Actions | [Submit Model](#) | [Comment/Bug](#)

Model | **Overview** | Math | Physical entities | Parameters | Curation

Create a submodel with selected elements

Deselect All

Model

Publication ID: [1831270](#) | **Submission Date:** 2005-09-13T12:31:08+00:00 | **Last Modification Date:** 2009-02-25T14:58:48+00:00 | **Creation Date:** 2005-02-08T18:28:27+00:00

Mathematical 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 degradation of cyclin |
| ☐ 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 | ☐ cyclin |
| | ☐ p-cyclin | ☐ total_cyclin | ☐ total_cdc2 |

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BIOMD0000000005 - Tyson1991_CellCycle_6var

SBML formats | Other formats | Actions | Submit Model Comment/Bug

Model Overview **Math** Physical entities Parameters

Reactions (2)

cdc2k phosphorylation $[cdc2k] \rightarrow [cdc2k-P];$

Math: $cell \times C2 \times k8notP$ (Detail)

Annotations: set #1 bqbiol:isVersionOf [Enzyme Nomenclature 2.7.11.1](#)
[Gene Ontology](#) protein amino acid phosphorylation

cdc2k dephosphorylation $[cdc2k-P] \rightarrow [cdc2k];$

Math: $cell \times CP \times k9$ (Detail)

Annotations: set #1 bqbiol:isVersionOf [Enzyme Nomenclature 3.1.3.16](#)
[Gene Ontology](#) protein amino acid dephosphorylation

Developed by BioModels Team of [Computational Neurobiology Group](#) in [European Bioinformatics Institute](#). | [Terms of Use](#) | [Contact Us](#)

MathML - Mozilla Firefox

http://www.ebi.ac.uk/biomodels-main/mathml.do?uri:

Reaction:

cdc2k dephosphorylation

rate law:

$cell * cdc2k-P * k9$

Compartments

Name	Size
cell	1.0

Species

Name	Compartment	Initial Amount	Initial Concentration
cdc2k-P	cell	0.75	

Parameters

Name	Value
k9	1000.0

Close

Done

zotero

BIOMD0000000005 - Tyson1991_CellCycle_6var



[SBML formats](#) |
 [Other formats](#) |
 [Actions](#) |
 [Submit Model Comment/Bug](#)

[Model](#) |
 [Overview](#) |
 [Math](#) |
 Physical entities |
 [Parameters](#) |
 [Curation](#)

cell	Spatial dimensions: 3 Compartment size: 1.0
cdc2k	Initial amount: 0.0
Compartment: cell	
Annotations:	set #1 bqbiol:isVersionOf UniProt CDC2_SCHPO
cdc2k-P	Initial amount: 0.75
Compartment: cell	
Annotations:	set #1 bqbiol:isVersionOf UniProt CDC2_SCHPO

BIOMD0000000005 - Tyson1991_CellCycle_6var



SBML formats

SBML L2 V1 (auto-generated)

SBML L2 V2 (auto-generated)

SBML L2 V3 (auto-generated)

SBML L2 V4 (curated)

Other formats (auto-generated)

Overview

Actions

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[Comment/Bug](#)

Publication ID: [1831270](#)

Proc Natl Acad Sci U S A. 1991; 88(16): 7328-32. [\[more\]](#)

Original Model: [BIOMD0000000005.xml](#)

Submitter: [Nicolas Le Novère](#)

Submission ID: MODEL6614644188

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

Last Modification Date: 10 Aug 2009 14:09:39 UTC

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

Encoders: [Bruce Shapiro](#)
[Vijayalakshmi Chelliah](#)

set #1

bqbiol:isVersionOf

bqmodel:is

[KEGG Pathway sce04111](#)

[Gene Ontology mitotic cell cycle](#)

[Taxonomy Fungi/Metazoa group](#)

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

Curated version of the model

Notes

This is a model from the article:
Modeling the cell division cycle: cdc2 and cyclin interactions.
Tyson JJ *Proc. Natl. Acad. Sci. U.S.A.* 1991; 88(16): 7328-32 [1831270](#).

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The proteins cdc2 and cyclin form a heterodimer (maturation promoting factor) that controls the major events of the cell cycle. A mathematical model for the interactions of cdc2 and cyclin is constructed. Simulation and analysis of the model show that the control system can operate in three modes: as a steady state with high maturation promoting factor activity, as a spontaneous oscillator, or as an excitable switch. We associate the steady state with metaphase arrest in unfertilized eggs, the spontaneous oscillations with rapid division cycles in early embryos, and the excitable switch with growth-controlled division cycles typical of nonembryonic cells.

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BIOMD0000000005 - Tyson1991_CellCycle_6var

SBML formats

Other formats (auto-generated)

Actions

Model

CellML

SciLab

XPP

BioPAX

VCell

PDF

Publication ID: 1831270

Model description

Division cycle: cdc2 and cyclin interactions

Department of Biology, Virginia Polytechnic Institute and State University

Original Model: [BIOMD0000000005.xml.orig](#)

Submitter: [Nicolas Le Novère](#)

Submission ID: MODEL6614644188

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

Last Modification Date: 10 Aug 2009 14:09:39 UTC

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

Encoders: [Bruce Shapiro](#)
[Vijayalakshmi Chelliah](#)

Model details

bqbiol:hasVersion [Reactome REACT_152](#)

set #1 bqbiol:isVersionOf [KEGG Pathway sce04111](#)
[Gene Ontology mitotic cell cycle](#)

bqmodel:is [Taxonomy Funqi/Metazoa group](#)

Notes

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```
<rdf:RDF
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  xmlns:bp="http://www.biopax.org/release/biopax-level2.owl#"
  xmlns:owl="http://www.w3.org/2002/07/owl#"
  xmlns="http://www.ebi.ac.uk/biomodels/biopax.owl#"
  xmlns:daml="http://www.daml.org/2001/03/daml-oi1#"
  xmlns:rdfs="http://www.w3.org/2000/01/rdf-schema#"
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  </owl:Ontology>
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    <bp:ID:G0:0005623</bp:ID>
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    <bp:CELLULAR-LOCATION>
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        <bp:XREF rdf:resource="#Gene_Ontology_G0_0005623"/>
        <bp:TERM:cell</bp:TERM>
      </bp:openControlledVocabulary>
    </bp:CELLULAR-LOCATION>
    <bp:PHYSICAL-ENTITY>
      <bp:protein rdf:ID="C2">
        <bp:XREF>
          <bp:unificationXref rdf:ID="UniProt_P04551">
            <bp:ID:P04551</bp:ID>
            <bp:DB:UniProt</bp:DB>
          </bp:unificationXref>
        </bp:XREF>
        <bp:NAME>cdc2k</bp:NAME>
      </bp:protein>
    </bp:PHYSICAL-ENTITY>
  </bp:physicalEntityParticipant>
  <bp:unificationXref rdf:ID="Gene_Ontology_G0_0006470">
    <bp:ID:G0:0006470</bp:ID>
    <bp:DB:Gene_Ontology</bp:DB>
  </bp:unificationXref>
  <bp:protein rdf:ID="CP">
    <bp:XREF rdf:resource="#UniProt_P04551"/>
    <bp:NAME>cdc2k-P</bp:NAME>
  </bp:protein>
  <bp:physicalEntityParticipant rdf:ID="Reaction5">
    <bp:STOICHIOMETRIC-COEFFICIENT>1.0</bp:STOICHIOMETRIC-COEFFICIENT>
    <bp:CELLULAR-LOCATION rdf:resource="#ce11"/>
    <bp:PHYSICAL-ENTITY>
```

SBML Model Report

Model name: "Tyson1991_CellCycle_6var"

9th February 2009

1 General Overview

This is a document in SBML Level 2 Version 3 format. This model was created by the following two authors: Bruce Shapiro¹ and Vijayalakshmi Chelliah² at February eighth 2005 at 6:28 p.m. and last modified at August 21st 2008 at 11:31 a.m. Table 1 gives an overview of the quantities of all components of this model.

Table 1: The SBML components in this model.
All components are described in more detail in the following sections.

Element	Quantity	Element	Quantity
compartment types	0	compartments	1
species types	0	species	9
events	0	constraints	0
reactions	9	function definitions	0
global parameters	0	unit definitions	5
rules	2	initial assignments	0

Model Notes

Cell Cycle Model; Tyson (1991, 6 variables)

Description

¹NASA Jet Propulsion Laboratory, bshapiro@jpl.nasa.gov
²EMBL-EBI, vij@ebi.ac.uk

Produced by SBML2LATEX



BIOMD0000000005 - Tyson1991_CellCycle_6var

SBML formats | Other formats (auto-generated) | Actions

Model Overview Math

View Bitmap Reaction Graph
View SVG Reaction Graph
View Dynamic Reaction Graph
View Model of Month

Publication ID: [1831270](#)

Proc Natl Acad Sci
Modeling the cell division cycle
Tyson JJ.
Department of Biology, Virginia Polytechnic Institute and State University

JWS Online Simulation
BioModels Online Simulation

Original Model: [BIOMD0000000005.xml.origin](#)

Submitter: [Nicolas Le Novère](#)

Submission ID: MODEL6614644188

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

Last Modification Date: 10 Aug 2009 14:09:39 UTC

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

Encoders: [Bruce Shapiro](#)
[Vijayalakshmi Chelliah](#)

This is a model from the article:
Modeling the cell division cycle: cdc2 and cyclin interactions.
Tyson JJ *Proc. Natl. Acad. Sci. U.S.A.* 1991; 88(16): 7328-32 [1831270](#).

Abstract:
The proteins cdc2 and cyclin form a heterodimer (maturation promoting factor) that controls the major events of the cell cycle. A number of the model show that the control system can operate in three modes: as a steady state with high maturation promoting factor activity, as a steady state with low maturation promoting factor activity, as a steady state with intermediate maturation promoting factor activity, as a steady state with high maturation promoting factor activity and metaphase arrest in unfertilized eggs, the spontaneous oscillations with rapid division cycles in early embryos, and the excitable system in late embryos.

This model originates from BioModels Database: A Database of Annotated Published Models. It is copyright (c) 2005-2010 The BioModels Consortium. For more information see the [terms of use](#).
To cite BioModels Database, please use [Le Novère N., Bornstein B., Broicher A., Courtot M., Donizelli M., Dharuri H., Li L., Sauro H., Schuster P., Shapiro B., Vajda S., Vajayalakshmi Chelliah, Curated, Published, Quantitative Kinetic Models of Biochemical and Cellular Systems Nucleic Acids Res. 34: D689-D691.](#)

http://www.ebi.ac.uk/biomodels-main/publ-model-tab.do?cmd=MODEL:SIMU

Model - Simulation

For doing an online simulation, please select the species below. After specifying the simulation time and print step, and then click **Submit** to submit simulation job to our research cluster.

Click **Cancel** to close the window.

☐ Species

☐ EmptySet

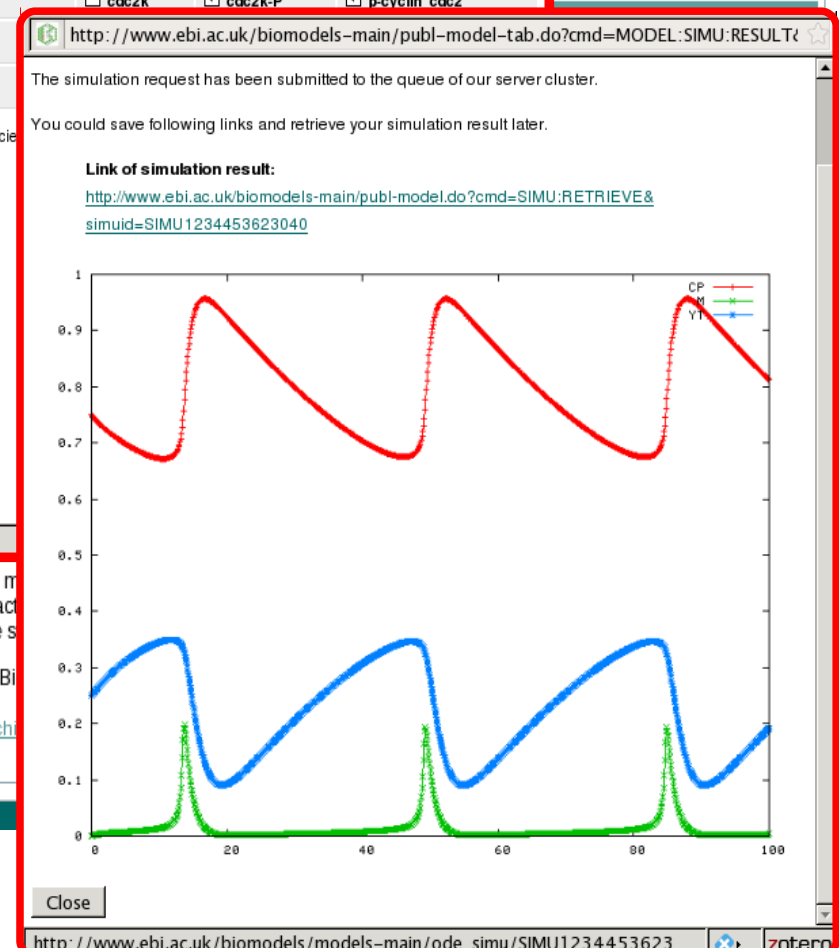
☐ p-cyclin_cdc2-p

☐ total_cdc2

☐ cdc2k ☒ cdc2k-P ☒ p-cyclin_cdc2

Simulation Time (use seconds):

Print step:



BIOMD0000000005 - Tyson1991_CellCycle_6var

SBML formats
Other formats (auto-generated)
Actions
[View Bitmap Reaction Graph](#)
[View SVG Reaction Graph](#)
[View Dynamic Reaction Graph](#)
[View Model of Month](#)
[JWS Online Simulation](#)
[BioModels Online Simulation](#)
[Submit Model](#)
[Comment/Bug](#)

Model
Overview
Math
Parameters
Reference Publication

Publication ID: [1831270](#)
Proc Natl Acad Sci
Modeling the cell di
Tyson JJ.
Department of Biology, Virginia Polytechnic Institute and State University, Blacksburg 240

Original Model: [BIOMD0000000005.xml.origin](#)
bqbiol:hasVersion [Reactome REACT_152](#)

Submitter: [Nicolas Le Novère](#)
set #1 bqbiol:isVersionOf [KEGG Pathway sce04111](#)
[Gene Ontology mitotic cell cycle](#)

Submission ID: MODEL6614644188
bqmodel:is [Taxonomy Funqi/Metazoa group](#)

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

Last Modification Date: 10 Aug 2009 14:09:39 UTC

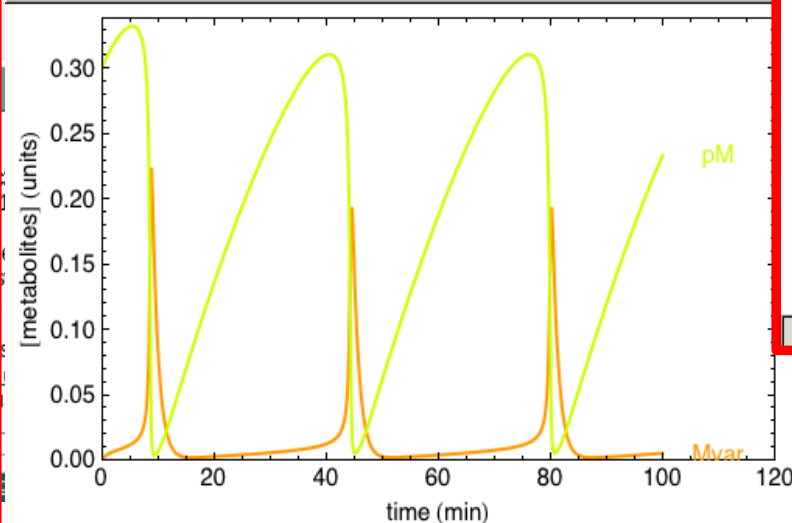
Creation Date: 08 Feb 2005 18:28:27 UTC
<http://jii.mib.ac.uk/webMathematica/Examples/PlotScript11.jsp?fun='JWS'>

Encoders: [Bruce Shapiro](#)
[Vijayalakshmi Chelliah](#)

This a model from the article:
Modeling the cell division cycle: cd
Tyson JJ *Proc. Natl. Acad. Sci. U.S.A.*
Abstract:
The proteins cdc2 and cyclin form a
of the model show that the control sys
metaphase arrest in unfertilized eggs

This model originates from BioMode
For more information see the [terms of use](#)
To cite BioModels Database, please
[Cited, Published, Quantitative Kinet](#)

[Computational Neurobiology Group in Eu](#)



Download the results in text or comma separated value format (e.g. for Excel import):

<http://jii.mib.ac.uk/biomodels/Tyson1991/index.html>

[Biomodels Home](#)
[Index](#)
[JWS Online](#)

Biomodels: BIOMD0000000005
Tyson1991
Powered by JWS Online

JWSApplet - ver 5.0.3 Tyson1991

	Parameter	Value
P1_v1	cell	1
P2_v1	k6	1
P3_v1	k8notP	1.0e6
P4_v1	k9	1000
P5_v1	k3	200
P6_v1	k5notP	0.
P7_v1	k1aa	0.015
P8_v1	k2	0.
P9_v1	k7	0.6
P10_v1	k4	180
P11_v1	k4prime	0.018
E1	EmptySet	0.
I1	C2[0]	0.
I2	CP[0]	1
I3	Mvar[0]	0.
I4	Y[0]	0.
I5	YP[0]	0.
I6	pM[0]	0.3

Evaluate Model

Sim State

Start Time End Time
0 100

☐ Rates ☒ Metabolites

Type	Output	Plot
M1	C2	☐
M2	CP	☐
M3	Mvar	☒
M4	Y	☐
M5	YP	☐
M6	pM	☒
F1	vRea...	☐
F2	vRea...	☐
F3	vRea...	☐
F4	vRea...	☐
F5	vRea...	☐
F6	vRea...	☐
F7	vRea...	☐
F8	vRea...	☐

Reset

Applet jiiApplet started

[Snoep J.L., Hucka M. \(2006\) BioModels Database: A Free, Centralized Database of](#)

BIOMD0000000005 - Tyson1991_CellCycle_6var

SBML formats

Other formats

Actions

Submit Model Comment/Bug

Model

Overview

Math

Physical entities

Parameters

Curation

Submodel1

Create a submodel with selected elements

Deselect All

Model

Publication ID: [1831270](#)

Submission Date: 2005-09-13T12:31:08+00:00

Last Modification Date: 2009-02-25T14:58:48+00:00

Creation Date: 2005-02-08T18:28:27+00:00

Mathematical 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 degradation of cyclin |
| ☐ 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 | ☐ cyclin |
| | ☐ p-cyclin | ☐ total_cyclin | ☐ total_cdc2 |

BIOMD0000000005 - Tyson1991_CellCycle_6var



SBML formats | Other formats | Actions | [Submit Model Comment/Bug](#)

Model Overview Math Physical entities Parameters Curation **Submodel1**

View the submodel in SBML

Save as

Reactions (2)

cdc2k phosphorylation [cdc2k] → [cdc2k-P];

cdc2k dephosphorylation [cdc2k-P] → [cdc2k];

Compartments

cell set #1 bqbiol:is [Gene Ontology cell](#)

Referred to as: cell

Species

cdc2k Initial amount: 0.0

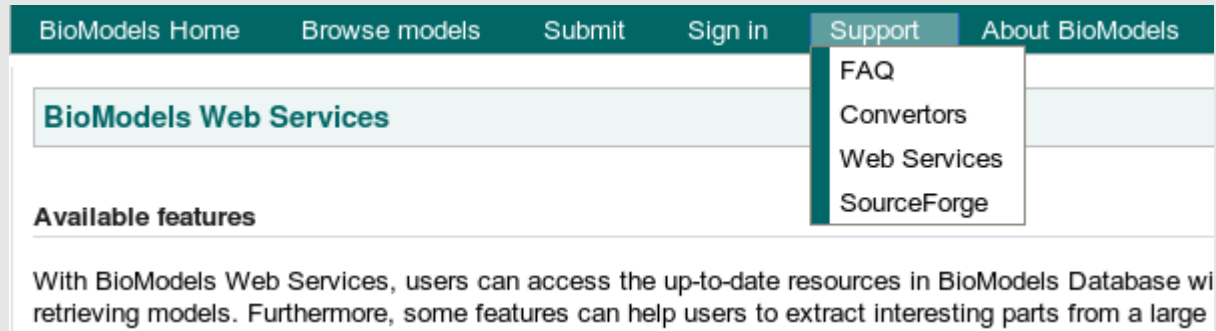
Compartment: cell

cdc2k-P Initial amount: 0.75

Compartment: cell

```
<?xml version="1.0" encoding="UTF-8"?>
<sbml xmlns="http://www.sbml.org/sbml/level2/version3" level="2" version="3">
  <model id="SUBMODEL1234454259626">
    <notes>
      <body xmlns="http://www.w3.org/1999/xhtml">
        <p>This is a sub-model automatically generated by BioModels Database. The generation
      </body>
    </notes>
    <listOfCompartments>
      <compartment metaid="_000002" id="cell" size="1">
        <annotation>
          <rdf:RDF xmlns:rdf="http://www.w3.org/1999/02/22-rdf-syntax-ns#" xmlns:dc="http://
            <rdf:Description rdf:about="#_000002">
              <bqbiol:is>
                <rdf:Bag>
                  <rdf:li rdf:resource="urn:miriam:obo:go:GO%3A0005623"/>
                </rdf:Bag>
              </bqbiol:is>
            </rdf:Description>
          </rdf:RDF>
        </annotation>
      </compartment>
    </listOfCompartments>
    <listOfSpecies>
      <species metaid="_000004" id="C2" name="cdc2k" compartment="cell" initialAmount="0">
        <annotation>
          <rdf:RDF xmlns:rdf="http://www.w3.org/1999/02/22-rdf-syntax-ns#" xmlns:dc="http://
            <rdf:Description rdf:about="#_000004">
              <bqbiol:isVersionOf>
                <rdf:Bag>
                  <rdf:li rdf:resource="urn:miriam:uniprot:P04551"/>
                </rdf:Bag>
              </bqbiol:isVersionOf>
            </rdf:Description>
          </rdf:RDF>
        </annotation>
      </species>
    </listOfSpecies>
  </model>
</sbml>
```

Software and Webservices



- Convertors: convertors for SBML to various formats, tools for graph generation and other related software
(<http://www.ebi.ac.uk/compneur-srv/sbml/convertors/SBMLConvertors.html>)
- Web Services: <http://www.ebi.ac.uk/biomodels-main/webservices>
Java library for accessing Biomodels DB, searching models, creating submodels, ...
- BiomodelsDB: <http://sourceforge.net/projects/biomodels/>
(latest code available on demand)

Nicolas
Le Novère



Michael
Hucka



Development

Camille Laibe



Nicolas Rodriguez



Curation

Lukas Endler



Vijayalakshmi
Chelliah



Melanie Stefan



In cooperation with:

Upinder Bhalla (DOQCS, NCBS, Bangalore)

Ion Moraru (the **Virtual Cell**, University of Connecticut Health Center)

Jacky Snoep (JWS Online, Stellenbosch (ZA) University, ZA)



Major Differences to the CellML Model Repository

- models need to be described in the peer reviewed literature
 - no public access to unpublished models
- models are only publicly available after the curation process
- two branches
 - curated: manually checked and annotated models
 - non curated: valid SBML, but neither checked nor annotated
- elements of curated models are manually annotated
- models, model elements and annotations are stored in a database
- models have unique perennial IDs, that can be used for retrieval

Minimal Information Requested in the Annotation of Biochemical Models (MIRIAM)*

- encoded in a machine readable format
 - SBML (<http://sbml.org>)
 - CellML for submission (<http://www.cellml.org>)
- clearly related to a peer reviewed reference publication containing a set of results
- model's structure must reflect the biological processes in the reference description
- the model must reproduce the results given in the reference description

*Nicolas Le Novère et al., *Nature Biotechnology*, **23**(12), 2005