# **SBML Model Report**

# Model name: "Tyson1991 - Cell Cycle 6 var"



May 6, 2016

#### 1 General Overview

This is a document in SBML Level 2 Version 4 format. This model was created by the following two authors: Bruce Shapiro<sup>1</sup> and Vijayalakshmi Chelliah<sup>2</sup> at February eighth 2005 at 6:28 p.m. and last time modified at May 16<sup>th</sup> 2013 at 2:40 p.m. Table 1 gives an overview of the quantities of all components of this model.

Table 1: Number of components in this model, which are described 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	0
rules	2	initial assignments	0

#### **Model 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. U.S.A. 1991; 88(16); 7328-32

Abstract:

<sup>&</sup>lt;sup>1</sup>NASA Jet Propulsion Laboratory, bshapiro@jpl.nasa.gov

 $<sup>^2</sup>$ EMBL-EBI, viji@ebi.ac.uk

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.

This model is hosted on BioModels Database and identified by: BIOMD0000000005.

To cite BioModels Database, please use: BioModels Database: An enhanced, curated and annotated resource for published quantitative kinetic models.

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#### 2 Unit Definitions

This is an overview of five unit definitions which are all predefined by SBML and not mentioned in the model.

#### 2.1 Unit substance

**Notes** Mole is the predefined SBML unit for substance.

**Definition** mol

#### 2.2 Unit volume

**Notes** Litre is the predefined SBML unit for volume.

**Definition** 1

#### 2.3 Unit area

**Notes** Square metre is the predefined SBML unit for area since SBML Level 2 Version 1.

**Definition** m<sup>2</sup>

#### 2.4 Unit length

**Notes** Metre is the predefined SBML unit for length since SBML Level 2 Version 1.

**Definition** m

## 2.5 Unit time

 $\mbox{\bf Notes}\,$  Second is the predefined SBML unit for time.

**Definition** s

# 3 Compartment

This model contains one compartment.

Table 2: Properties of all compartments.

Id	Name	SBO	Spatial Dimensions	Size	Unit	Constant	Outside
cell			3	1	litre	Ø	

# **3.1 Compartment** cell

This is a three dimensional compartment with a constant size of one litre.

# 4 Species

This model contains nine species. The boundary condition of three of these species is set to true so that these species' amount cannot be changed by any reaction. Section 7 provides further details and the derived rates of change of each species.

Table 3: Properties of each species.

Id	Name	Compartment	Derived Unit	Constant	Boundary Condi- tion
EmptySet		cell	$\text{mol} \cdot 1^{-1}$		$ \overline{\mathbf{Z}} $
C2	cdc2k	cell	$\text{mol} \cdot l^{-1}$		
CP	cdc2k-P	cell	$\text{mol} \cdot l^{-1}$		$\Box$
M	p-cyclin_cdc2	cell	$\operatorname{mol} \cdot 1^{-1}$		
pМ	p-cyclin_cdc2-p	cell	$\text{mol} \cdot l^{-1}$		$\Box$
Y	cyclin	cell	$\operatorname{mol} \cdot 1^{-1}$		$\Box$
YP	p-cyclin	cell	$\mathrm{mol}\cdot\mathrm{l}^{-1}$		
YT	total_cyclin	cell	$\operatorname{mol} \cdot 1^{-1}$		
CT	total_cdc2	cell	$\text{mol} \cdot l^{-1}$		

# 5 Rules

This is an overview of two rules.

## 5.1 Rule YT

Rule YT is an assignment rule for species YT:

$$YT = [Y] + [YP] + [M] + [pM]$$
 (1)

Derived unit  $mol \cdot l^{-1}$ 

# 5.2 Rule CT

Rule CT is an assignment rule for species CT:

$$CT = [C2] + [CP] + [M] + [pM]$$
 (2)

Derived unit  $mol \cdot l^{-1}$ 

# 6 Reactions

This model contains nine reactions. All reactions are listed in the following table and are subsequently described in detail. If a reaction is affected by a modifier, the identifier of this species is written above the reaction arrow.

Table 4: Overview of all reactions

Nº	Id	Name	Reaction Equation	SBO
1	Reaction1	cyclin_cdc2k dissociation	$M \longrightarrow C2 + YP$	
2	Reaction2	cdc2k phosphorylation	$C2 \longrightarrow CP$	
3	Reaction3	cdc2k dephosphorylation	$CP \longrightarrow C2$	
4	Reaction4	cyclin cdc2k-p association	$CP + Y \longrightarrow pM$	
5	Reaction5	deactivation of cdc2 kinase	$\mathbf{M} \longrightarrow \mathbf{p}\mathbf{M}$	
6	Reaction6	cyclin biosynthesis	$EmptySet \longrightarrow Y$	
7	Reaction7	default degradation of cyclin	$Y \longrightarrow EmptySet$	
8	Reaction8	cdc2 kinase triggered degration of cyclin	$YP \longrightarrow EmptySet$	
9	Reaction9	activation of cdc2 kinase	$pM \xrightarrow{CT} M$	

## **6.1 Reaction** Reaction1

This is an irreversible reaction of one reactant forming two products.

Name cyclin\_cdc2k dissociation

## **Reaction equation**

$$M \longrightarrow C2 + YP$$
 (3)

#### Reactant

Table 5: Properties of each reactant.

Id	Name	SBO
M	p-cyclin_cdc2	

#### **Products**

Table 6: Properties of each product.

Id	Name	SBO
C2	cdc2k	
ΥP	p-cyclin	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_1 = \text{vol}(\text{cell}) \cdot \text{k6} \cdot [\text{M}] \tag{4}$$

Table 7: Properties of each parameter.

k6		1.0	<b>I</b>
Id	Name	SBO Value Unit	Constant

#### **6.2 Reaction Reaction2**

This is an irreversible reaction of one reactant forming one product.

Name cdc2k phosphorylation

## **Reaction equation**

$$C2 \longrightarrow CP$$
 (5)

#### Reactant

Table 8: Properties of each reactant.

Id	Name	SBO
C2	cdc2k	

#### **Product**

Table 9: Properties of each product.

Id	Name	SBO
CP	cdc2k-P	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_2 = \text{vol}\left(\text{cell}\right) \cdot [\text{C2}] \cdot \text{k8notP}$$
 (6)

Table 10: Properties of each parameter.

Id	Name	SBO	Value	Unit	Constant
k8notP			1000000.0		

## **6.3 Reaction Reaction3**

This is an irreversible reaction of one reactant forming one product.

Name cdc2k dephosphorylation

# **Reaction equation**

$$CP \longrightarrow C2$$
 (7)

# Reactant

Table 11: Properties of each reactant.

Id	Name	SBO
CP	cdc2k-P	

#### **Product**

Table 12: Properties of each product.

Id	Name	SBO
C2	cdc2k	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_3 = \text{vol}(\text{cell}) \cdot [\text{CP}] \cdot \text{k9}$$
 (8)

Table 13: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k9		1000.0	

#### **6.4 Reaction Reaction4**

This is an irreversible reaction of two reactants forming one product.

Name cyclin cdc2k-p association

## **Reaction equation**

$$CP + Y \longrightarrow pM$$
 (9)

#### **Reactants**

Table 14: Properties of each reactant.

Id	Name	SBO
CP	cdc2k-P	
Y	cyclin	

## **Product**

Table 15: Properties of each product.

Id	Name	SBO
pM	p-cyclin_cdc2-p	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_4 = \text{vol}(\text{cell}) \cdot [\text{CP}] \cdot \text{k3} \cdot [\text{Y}]$$
 (10)

Table 16: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k3		200.0	

#### 6.5 Reaction Reaction5

This is an irreversible reaction of one reactant forming one product.

Name deactivation of cdc2 kinase

## **Reaction equation**

$$\mathbf{M} \longrightarrow \mathbf{pM}$$
 (11)

#### Reactant

Table 17: Properties of each reactant.

Id	Name	SBO
M	p-cyclin_cdc2	

#### **Product**

Table 18: Properties of each product.

Id	Name	SBO
pМ	p-cyclin_cdc2-p	

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#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_5 = \text{vol}(\text{cell}) \cdot \text{k5notP} \cdot [M] \tag{12}$$

Table 19: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k5notP		0.0	

#### 6.6 Reaction Reaction6

This is an irreversible reaction of one reactant forming one product.

Name cyclin biosynthesis

## **Reaction equation**

$$EmptySet \longrightarrow Y \tag{13}$$

#### Reactant

Table 20: Properties of each reactant.

Id	Name	SBO
EmptySet		

#### **Product**

Table 21: Properties of each product.

Id	Name	SBO
Y	cyclin	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_6 = \text{vol}\left(\text{cell}\right) \cdot \text{k1aa}$$
 (14)

Table 22: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k1aa		0.015	$\overline{\hspace{1cm}}$

#### **6.7 Reaction** Reaction 7

This is an irreversible reaction of one reactant forming one product.

Name default degradation of cyclin

## **Reaction equation**

$$Y \longrightarrow EmptySet$$
 (15)

#### Reactant

Table 23: Properties of each reactant.

Id	Name	SBO
Y	cyclin	

#### **Product**

Table 24: Properties of each product.

Id	Name	SBO
EmptySet		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_7 = \text{vol}(\text{cell}) \cdot \text{k2} \cdot [Y] \tag{16}$$

Table 25: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k2		0.0	

#### 6.8 Reaction Reaction8

This is an irreversible reaction of one reactant forming one product.

Name cdc2 kinase triggered degration of cyclin

## **Reaction equation**

$$YP \longrightarrow EmptySet$$
 (17)

#### Reactant

Table 26: Properties of each reactant.

Id	Name	SBO
ΥP	p-cyclin	

#### **Product**

Table 27: Properties of each product.

Id	Name	SBO
EmptySet		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_8 = \text{vol}\left(\text{cell}\right) \cdot \text{k7} \cdot [\text{YP}] \tag{18}$$

Table 28: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k7		0.6	

#### **6.9 Reaction** Reaction9

This is an irreversible reaction of one reactant forming one product influenced by one modifier.

Name activation of cdc2 kinase

## **Reaction equation**

$$pM \xrightarrow{CT} M \tag{19}$$

#### Reactant

Table 29: Properties of each reactant.

Id	Name	SBO
pM	p-cyclin_cdc2-p	

#### **Modifier**

Table 30: Properties of each modifier.

Id	Name	SBO
CT	total_cdc2	

#### **Product**

Table 31: Properties of each product.

Id	Name	SBO
М	p-cyclin_cdc2	

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_9 = \text{vol}(\text{cell}) \cdot [\text{pM}] \cdot \left(\text{k4prime} + \text{k4} \cdot \left(\frac{[\text{M}]}{[\text{CT}]}\right)^2\right)$$
 (20)

Table 32: Properties of each parameter.

Id	Name	SBO	Value	Unit	Constant
k4			180.000		$ \overline{\checkmark} $
k4prime			0.018		$\overline{\mathbf{Z}}$

# 7 Derived Rate Equations

When interpreted as an ordinary differential equation framework, this model implies the following set of equations for the rates of change of each species.

Identifiers for kinetic laws highlighted in gray cannot be verified to evaluate to units of SBML substance per time. As a result, some SBML interpreters may not be able to verify the consistency of the units on quantities in the model. Please check if

- parameters without an unit definition are involved or
- volume correction is necessary because the hasOnlySubstanceUnits flag may be set to false and spacialDimensions > 0 for certain species.

#### 7.1 Species EmptySet

#### Initial amount 0 mol

This species takes part in three reactions (as a reactant in Reaction6 and as a product in Reaction7, Reaction8), which do not influence its rate of change because this species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{EmptySet} = 0\tag{21}$$

# 7.2 Species C2

Name cdc2k

Initial amount 0 mol

This species takes part in three reactions (as a reactant in Reaction2 and as a product in Reaction1, Reaction3).

$$\frac{d}{dt}C2 = |v_1| + |v_3| - |v_2| \tag{22}$$

#### 7.3 Species CP

Name cdc2k-P

Initial amount 0.75 mol

This species takes part in three reactions (as a reactant in Reaction3, Reaction4 and as a product in Reaction2).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{CP} = |v_2| - |v_3| - |v_4| \tag{23}$$

#### 7.4 Species M

Name p-cyclin\_cdc2

Initial amount 0 mol

This species takes part in three reactions (as a reactant in Reaction1, Reaction5 and as a product in Reaction9).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathbf{M} = |\mathbf{v}_9| - |\mathbf{v}_1| - |\mathbf{v}_5| \tag{24}$$

# 7.5 Species pM

Name p-cyclin\_cdc2-p

Initial amount 0.25 mol

This species takes part in three reactions (as a reactant in Reaction9 and as a product in Reaction4, Reaction5).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{pM} = |v_4| + |v_5| - |v_9| \tag{25}$$

#### 7.6 Species Y

Name cyclin

Initial amount 0 mol

This species takes part in three reactions (as a reactant in Reaction4, Reaction7 and as a product in Reaction6).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathbf{Y} = |v_6| - |v_4| - |v_7| \tag{26}$$

## 7.7 Species YP

Name p-cyclin

Initial amount 0 mol

This species takes part in two reactions (as a reactant in Reaction8 and as a product in Reaction1).

$$\frac{\mathrm{d}}{\mathrm{d}t} \mathbf{Y} \mathbf{P} = |v_1| - |v_8| \tag{27}$$

## 7.8 Species YT

Name total\_cyclin

Involved in rule YT

One rule determines the species' quantity.

## 7.9 Species CT

Name total\_cdc2

Involved in rule CT

This species takes part in one reaction (as a modifier in Reaction9). Not this but one rule determines the species' quantity because this species is on the boundary of the reaction system.

 $\mathfrak{BML2}^{lAT}$ EX was developed by Andreas Dräger<sup>a</sup>, Hannes Planatscher<sup>a</sup>, Dieudonné M Wouamba<sup>a</sup>, Adrian Schröder<sup>a</sup>, Michael Hucka<sup>b</sup>, Lukas Endler<sup>c</sup>, Martin Golebiewski<sup>d</sup> and Andreas Zell<sup>a</sup>. Please see http://www.ra.cs.uni-tuebingen.de/software/SBML2LaTeX for more information.

<sup>&</sup>lt;sup>a</sup>Center for Bioinformatics Tübingen (ZBIT), Germany

<sup>&</sup>lt;sup>b</sup>California Institute of Technology, Beckman Institute BNMC, Pasadena, United States

<sup>&</sup>lt;sup>c</sup>European Bioinformatics Institute, Wellcome Trust Genome Campus, Hinxton, United Kingdom

<sup>&</sup>lt;sup>d</sup>EML Research gGmbH, Heidelberg, Germany