

## SBML Model Report

# Model name: “Chen2000 - Budding yeast cell cycle”



May 17, 2018

## 1 General Overview

This is a document in SBML Level 2 Version 4 format. This model was created by the following two authors: Matthew Grant Roberts<sup>1</sup> and Catherine Lloyd<sup>2</sup> at June 25<sup>th</sup> 2010 at 12:06 a. m. and last time modified at February 20<sup>th</sup> 2018 at 9:06 a. m. Table 1 provides 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	0
events	4	constraints	0
reactions	0	function definitions	0
global parameters	102	unit definitions	1
rules	29	initial assignments	0

## Model Notes

This a model from the article:

**Kinetic analysis of a molecular model of the budding yeast cell cycle.**

Chen KC, Csikasz-Nagy A, Gyorffy B, Val J, Novak B, Tyson JJ. Mol Biol Cell 2000 Jan;11(1):369-91 [10637314](#) ,

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**Abstract:**

The molecular machinery of cell cycle control is known in more detail for budding yeast, *Saccharomyces cerevisiae*, than for any other eukaryotic organism. In recent years, many elegant experiments on budding yeast have dissected the roles of cyclin molecules (Cln1-3 and Clb1-6) in coordinating the events of DNA synthesis, bud emergence, spindle formation, nuclear division, and cell separation. These experimental clues suggest a mechanism for the principal molecular interactions controlling cyclin synthesis and degradation. Using standard techniques of biochemical kinetics, we convert the mechanism into a set of differential equations, which describe the time courses of three major classes of cyclin-dependent kinase activities. Model in hand, we examine the molecular events controlling „Start,, (the commitment step to a new round of chromosome replication, bud formation, and mitosis) and „Finish,, (the transition from metaphase to anaphase, when sister chromatids are pulled apart and the bud separates from the mother cell) in wild-type cells and 50 mutants. The model accounts for many details of the physiology, biochemistry, and genetics of cell cycle control in budding yeast.

This model was taken from the [CellML repository](#) and automatically converted to SBML. The original model was: [Chen KC, Csikasz-Nagy A, Gyorffy B, Val J, Novak B, Tyson JJ. \(2000\) - version=1.0](#)

The original CellML model was created by:

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To cite BioModels Database, please use: [Li C, Donizelli M, Rodriguez N, Dharuri H, Endler L, Chelliah V, Li L, He E, Henry A, Stefan MI, Snoep JL, Hucka M, Le Novre N, Laibe C \(2010\) BioModels Database: An enhanced, curated and annotated resource for published quantitative kinetic models. BMC Syst Biol., 4:92.](#)

## 2 Unit Definitions

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

## 2.1 Unit time

**Name** time

**Definition** 60 s

## 2.2 Unit substance

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

**Definition** mol

## 2.3 Unit volume

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

**Definition** l

## 2.4 Unit area

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

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

## 2.5 Unit length

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

**Definition** m

# 3 Compartment

This model contains one compartment.

Table 2: Properties of all compartments.

Id	Name	SBO	Spatial Dimensions	Size	Unit	Constant	Outside
COMpartment	Yeast Cell		3	1	litre	<input checked="" type="checkbox"/>	

## 3.1 Compartment COMpartment

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

**Name** Yeast Cell

## 4 Parameters

This model contains 102 global parameters.

Table 3: Properties of each parameter.

Id	Name	SBO	Value	Unit	Constant
Cln2	Cln2		0.008		<input type="checkbox"/>
ks_n2	ks_n2_prime		0.000		<input checked="" type="checkbox"/>
ks_n2_	ks_n2_2prime		0.050		<input checked="" type="checkbox"/>
kd_n2	kd_n2		0.100		<input checked="" type="checkbox"/>
Clb2_T	Clb2_T		0.234		<input type="checkbox"/>
Hct1_T	Hct1_T		1.000		<input checked="" type="checkbox"/>
ks_b2	ks_b2_prime		0.002		<input checked="" type="checkbox"/>
ks_b2_	ks_b2_2prime		0.050		<input checked="" type="checkbox"/>
kd_b2	kd_b2_prime		0.010		<input checked="" type="checkbox"/>
kd_b2_	kd_b2_2prime		2.000		<input checked="" type="checkbox"/>
kd_b2_	kd_b2_3prime		0.050		<input checked="" type="checkbox"/>
Vd_b2	Vd_b2		2.023		<input type="checkbox"/>
Clb2	Clb2		0.155		<input type="checkbox"/>
Clb5	Clb5		0.041		<input type="checkbox"/>
Sic1	Sic1		0.023		<input type="checkbox"/>
Clb5_T	Clb5_T		0.061		<input type="checkbox"/>
ks_b5	ks_b5_prime		0.006		<input checked="" type="checkbox"/>
ks_b5_	ks_b5_2prime		0.020		<input checked="" type="checkbox"/>
kd_b5	kd_b5_prime		0.100		<input checked="" type="checkbox"/>
kd_b5_	kd_b5_2prime		0.250		<input checked="" type="checkbox"/>
Vd_b5	Vd_b5		0.271		<input type="checkbox"/>
Bck2	Bck2		0.002		<input type="checkbox"/>
Bck2_0	Bck2_0		0.003		<input checked="" type="checkbox"/>
Cln3	Cln3		0.002		<input type="checkbox"/>
Jn3	Jn3		6.000		<input checked="" type="checkbox"/>
Dn3	Dn3		1.000		<input checked="" type="checkbox"/>
Cln3_max	Cln3_max		0.020		<input checked="" type="checkbox"/>
Sic1_T	Sic1_T		0.123		<input type="checkbox"/>
ks_c1	ks_c1		0.020		<input checked="" type="checkbox"/>
ks_c1_	ks_c1_2prime		0.100		<input checked="" type="checkbox"/>
Clb2_Sic1	Clb2_Sic1		0.079		<input type="checkbox"/>
kas_b2	kas_b2		50.000		<input checked="" type="checkbox"/>
kdi_b2	kdi_b2		0.050		<input checked="" type="checkbox"/>
Clb5_Sic1	Clb5_Sic1		0.021		<input type="checkbox"/>
kas_b5	kas_b5		50.000		<input checked="" type="checkbox"/>
kdi_b5	kdi_b5		0.050		<input checked="" type="checkbox"/>

Id	Name	SBO	Value	Unit	Constant
Vd2_c1	Vd2_c1		0.031		<input type="checkbox"/>
kd2_c1	kd2_c1		0.300		<input checked="" type="checkbox"/>
epsilon_c1_n3	epsilon_c1_n3		20.000		<input checked="" type="checkbox"/>
epsilon_c1_k2	epsilon_c1_k2		2.000		<input checked="" type="checkbox"/>
epsilon_c1_b5	epsilon_c1_b5		1.000		<input checked="" type="checkbox"/>
epsilon_c1_b2	epsilon_c1_b2		0.067		<input checked="" type="checkbox"/>
Cdc20_T	Cdc20_T		0.833		<input type="checkbox"/>
ks_20	ks_20_prime		0.005		<input checked="" type="checkbox"/>
ks_20_	ks_20_2prime		0.060		<input checked="" type="checkbox"/>
Cdc20	Cdc20		0.685		<input type="checkbox"/>
ka_20	ka_20		1.000		<input checked="" type="checkbox"/>
ki_20	ki_20_prime		0.100		<input checked="" type="checkbox"/>
ki_20_	ki_20_2prime		10.000		<input checked="" type="checkbox"/>
Vi_20	Vi_20		0.100		<input type="checkbox"/>
Hct1	Hct1		0.995		<input type="checkbox"/>
ka_t1	ka_t1_prime		0.040		<input checked="" type="checkbox"/>
ka_t1_	ka_t1_2prime		2.000		<input checked="" type="checkbox"/>
ki_t1	ki_t1_prime		0.000		<input checked="" type="checkbox"/>
ki_t1_	ki_t1_2prime		0.640		<input checked="" type="checkbox"/>
Vi_t1	Vi_t1		0.119		<input type="checkbox"/>
Ji_t1	Ji_t1		0.050		<input checked="" type="checkbox"/>
Ja_t1	Ja_t1		0.050		<input checked="" type="checkbox"/>
epsilon_i_t1-_n2	epsilon_i_t1_n2		1.000		<input checked="" type="checkbox"/>
epsilon_i_t1-_b5	epsilon_i_t1_b5		0.500		<input checked="" type="checkbox"/>
epsilon_i_t1-_b2	epsilon_i_t1_b2		1.000		<input checked="" type="checkbox"/>
mass	mass		0.661		<input type="checkbox"/>
mu	mu		0.006		<input checked="" type="checkbox"/>
ORI	ORI		0.000		<input type="checkbox"/>
ks_ori	ks_ori		2.000		<input checked="" type="checkbox"/>
kd_ori	kd_ori		0.060		<input checked="" type="checkbox"/>
epsilon_ori-_b2	epsilon_ori_b2		0.400		<input checked="" type="checkbox"/>
BUD	BUD		0.000		<input type="checkbox"/>
ks_bud	ks_bud		0.300		<input checked="" type="checkbox"/>
kd_bud	kd_bud		0.060		<input checked="" type="checkbox"/>
epsilon_bud-_b5	epsilon_bud_b5		1.000		<input checked="" type="checkbox"/>
SPN	SPN		0.000		<input type="checkbox"/>
ks_spn	ks_spn		0.080		<input checked="" type="checkbox"/>

Id	Name	SBO	Value	Unit	Constant
kd_spn	kd_spn		0.060		<input checked="" type="checkbox"/>
J_spn	J_spn		0.200		<input checked="" type="checkbox"/>
SBF	SBF		0.003		<input type="checkbox"/>
ka_sbf	ka_sbf		1.000		<input checked="" type="checkbox"/>
ki_sbf	ki_sbf_prime		0.500		<input checked="" type="checkbox"/>
ki_sbf_	ki_sbf_2prime		6.000		<input checked="" type="checkbox"/>
Va_sbf	Va_sbf		0.311		<input type="checkbox"/>
Ji_sbf	Ji_sbf		0.010		<input checked="" type="checkbox"/>
Ja_sbf	Ja_sbf		0.010		<input checked="" type="checkbox"/>
epsilonsbf-n3	epsilonsbf_n3		75.000		<input checked="" type="checkbox"/>
epsilonsbf-b5	epsilonsbf_b5		0.500		<input checked="" type="checkbox"/>
MBF	MBF		0.003		<input type="checkbox"/>
Mcm1	Mcm1		0.513		<input type="checkbox"/>
ka_mcm	ka_mcm		1.000		<input checked="" type="checkbox"/>
ki_mcm	ki_mcm		0.150		<input checked="" type="checkbox"/>
Ji_mcm	Ji_mcm		1.000		<input checked="" type="checkbox"/>
Ja_mcm	Ja_mcm		1.000		<input checked="" type="checkbox"/>
Swi5	Swi5		0.923		<input type="checkbox"/>
ka_swi	ka_swi		1.000		<input checked="" type="checkbox"/>
ki_swi	ki_swi_prime		0.300		<input checked="" type="checkbox"/>
ki_swi_	ki_swi_2prime		0.200		<input checked="" type="checkbox"/>
Ji_swi	Ji_swi		0.100		<input checked="" type="checkbox"/>
Ja_swi	Ja_swi		0.100		<input checked="" type="checkbox"/>
kd1_c1	kd1_c1		0.010		<input checked="" type="checkbox"/>
kd_20	kd_20		0.080		<input checked="" type="checkbox"/>
Jd2_c1	Jd2_c1		0.050		<input checked="" type="checkbox"/>
END_M	END_M		2000.000		<input type="checkbox"/>
START_S	START_S		1000.000		<input type="checkbox"/>
D	D		145.632		<input type="checkbox"/>

## 5 Rules

This is an overview of 29 rules.

### 5.1 Rule Vd\_b2

Rule Vd\_b2 is an assignment rule for parameter Vd\_b2:

$$Vd\_b2 = kd\_b2 \cdot (Hct1\_T - Hct1) + kd\_b2\_ \cdot Hct1 + kd\_b2\_ \cdot Cdc20 \quad (1)$$

## 5.2 Rule C1b2

Rule C1b2 is an assignment rule for parameter C1b2:

$$C1b2 = C1b2\_T - C1b2\_Sic1 \quad (2)$$

## 5.3 Rule C1b5

Rule C1b5 is an assignment rule for parameter C1b5:

$$C1b5 = C1b5\_T - C1b5\_Sic1 \quad (3)$$

## 5.4 Rule Sic1

Rule Sic1 is an assignment rule for parameter Sic1:

$$Sic1 = Sic1\_T - (C1b2\_Sic1 + C1b5\_Sic1) \quad (4)$$

## 5.5 Rule Vd\_b5

Rule Vd\_b5 is an assignment rule for parameter Vd\_b5:

$$Vd\_b5 = kd\_b5 + kd\_b5\_ \cdot Cdc20 \quad (5)$$

## 5.6 Rule Bck2

Rule Bck2 is an assignment rule for parameter Bck2:

$$Bck2 = Bck2\_0 \cdot mass \quad (6)$$

## 5.7 Rule Cln3

Rule Cln3 is an assignment rule for parameter Cln3:

$$Cln3 = \frac{Cln3\_max \cdot Dn3 \cdot mass}{Jn3 + Dn3 \cdot mass} \quad (7)$$

## 5.8 Rule Va\_sbf

Rule Va\_sbf is an assignment rule for parameter Va\_sbf:

$$Va\_sbf = ka\_sbf \cdot (Cln2 + \epsilon_{sbf\_n3} \cdot (Cln3 + Bck2) + \epsilon_{sbf\_b5} \cdot C1b5) \quad (8)$$

## 5.9 Rule Vd2\_c1

Rule Vd2\_c1 is an assignment rule for parameter Vd2\_c1:

$$Vd2\_c1 = kd2\_c1 \cdot (\epsilon_{c1\_n3} \cdot Cln3 + \epsilon_{c1\_k2} \cdot Bck2 + Cln2 + \epsilon_{c1\_b5} \cdot C1b5 + \epsilon_{c1\_b2} \cdot C1b2) \quad (9)$$

### 5.10 Rule $Vi\_20$

Rule  $Vi\_20$  is an assignment rule for parameter  $Vi\_20$ :

$$Vi\_20 = \begin{cases} 10 & \text{if } (time \geq START\_S) \wedge (time < END\_M + 12) \\ 10 - 9.9 \cdot \frac{time - END\_M}{12} & \text{if } (time \geq END\_M) \wedge (time < END\_M + 12) \\ 0.1 & \text{otherwise} \end{cases} \quad (10)$$

### 5.11 Rule $Vi\_t1$

Rule  $Vi\_t1$  is an assignment rule for parameter  $Vi\_t1$ :

$$Vi\_t1 = ki\_t1 + ki\_t1\_ \cdot (Cln3 + \epsilon_{t1\_n2} \cdot Cln2 + \epsilon_{t1\_b5} \cdot Clb5 + \epsilon_{t1\_b2} \cdot Clb2) \quad (11)$$

### 5.12 Rule $SBF$

Rule  $SBF$  is an assignment rule for parameter  $SBF$ :

$$SBF = \frac{2 \cdot V}{ki\_sbf + ki\_sbf\_ \cdot Clb2 + Va\_sbf \cdot Ji\_sbf + (ki\_sbf + ki\_sbf\_ \cdot Clb2) \cdot Ja\_sbf - Va\_sbf + ((ki\_sbf + ki\_sbf\_ \cdot Clb2) \cdot Ja\_sbf - Va\_sbf)} \quad (12)$$

### 5.13 Rule $MBF$

Rule  $MBF$  is an assignment rule for parameter  $MBF$ :

$$MBF = SBF \quad (13)$$

### 5.14 Rule $Mcm1$

Rule  $Mcm1$  is an assignment rule for parameter  $Mcm1$ :

$$Mcm1 = \frac{2 \cdot ka\_mcm \cdot Clb2}{ki\_mcm + ka\_mcm \cdot Clb2 \cdot Ji\_mcm + ki\_mcm \cdot Ja\_mcm - ka\_mcm \cdot Clb2 + ((ki\_mcm + ka\_mcm \cdot Clb2 \cdot Ji\_mcm) \cdot Ja\_mcm - ka\_mcm \cdot Clb2)} \quad (14)$$

### 5.15 Rule $Swi5$

Rule  $Swi5$  is an assignment rule for parameter  $Swi5$ :

$$Swi5 = \frac{ki\_swi + ki\_swi\_ \cdot Clb2 + ka\_swi \cdot Cdc20 \cdot Ji\_swi + (ki\_swi + ki\_swi\_ \cdot Clb2) \cdot Ja\_swi - ka\_swi \cdot Cdc20 + ((ki\_swi + ki\_swi\_ \cdot Clb2) \cdot Ja\_swi - ka\_swi \cdot Cdc20)}{2 \cdot V} \quad (15)$$



### 5.16 Rule D

Rule D is an assignment rule for parameter D:

$$D = \frac{1.026}{\mu} - 32 \quad (16)$$

### 5.17 Rule Cln2

Rule Cln2 is a rate rule for parameter Cln2:

$$\frac{d}{dt}Cln2 = mass \cdot (ks_{n2} + ks_{n2\_} \cdot SBF) - kd_{n2} \cdot Cln2 \quad (17)$$

### 5.18 Rule Clb2\_T

Rule Clb2\_T is a rate rule for parameter Clb2\_T:

$$\frac{d}{dt}Clb2\_T = mass \cdot (ks_{b2} + ks_{b2\_} \cdot Mcm1) - Vd_{b2} \cdot Clb2\_T \quad (18)$$

### 5.19 Rule Clb5\_T

Rule Clb5\_T is a rate rule for parameter Clb5\_T:

$$\frac{d}{dt}Clb5\_T = mass \cdot (ks_{b5} + ks_{b5\_} \cdot MBF) - Vd_{b5} \cdot Clb5\_T \quad (19)$$

### 5.20 Rule Sic1\_T

Rule Sic1\_T is a rate rule for parameter Sic1\_T:

$$\frac{d}{dt}Sic1\_T = ks_{c1} + ks_{c1\_} \cdot Swi5 - Sic1\_T \cdot \left( kd1_{c1} + \frac{Vd2_{c1}}{Jd2_{c1} + Sic1\_T} \right) \quad (20)$$

### 5.21 Rule Clb2\_Sic1

Rule Clb2\_Sic1 is a rate rule for parameter Clb2\_Sic1:

$$\begin{aligned} \frac{d}{dt}Clb2\_Sic1 &= kas_{b2} \cdot Clb2 \cdot Sic1 - Clb2\_Sic1 \\ &\cdot \left( kdi_{b2} + Vd_{b2} + kd1_{c1} + \frac{Vd2_{c1}}{Jd2_{c1} + Sic1\_T} \right) \end{aligned} \quad (21)$$

### 5.22 Rule Clb5\_Sic1

Rule Clb5\_Sic1 is a rate rule for parameter Clb5\_Sic1:

$$\begin{aligned} \frac{d}{dt}Clb5\_Sic1 &= kas_{b5} \cdot Clb5 \cdot Sic1 - Clb5\_Sic1 \\ &\cdot \left( kdi_{b5} + Vd_{b5} + kd1_{c1} + \frac{Vd2_{c1}}{Jd2_{c1} + Sic1\_T} \right) \end{aligned} \quad (22)$$

### 5.23 Rule Cdc20\_T

Rule Cdc20\_T is a rate rule for parameter Cdc20\_T:

$$\frac{d}{dt}Cdc20\_T = ks\_20 + ks\_20\_ \cdot Clb2 - kd\_20 \cdot Cdc20\_T \quad (23)$$

### 5.24 Rule Cdc20

Rule Cdc20 is a rate rule for parameter Cdc20:

$$\frac{d}{dt}Cdc20 = ka\_20 \cdot (Cdc20\_T - Cdc20) - Cdc20 \cdot (Vi\_20 + kd\_20) \quad (24)$$

### 5.25 Rule Hct1

Rule Hct1 is a rate rule for parameter Hct1:

$$\frac{d}{dt}Hct1 = \frac{(ka\_t1 + ka\_t1\_ \cdot Cdc20) \cdot (Hct1\_T - Hct1)}{Ja\_t1 + Hct1\_T - Hct1} - \frac{Vi\_t1 \cdot Hct1}{Ji\_t1 + Hct1} \quad (25)$$

### 5.26 Rule mass

Rule mass is a rate rule for parameter mass:

$$\frac{d}{dt}mass = mu \cdot mass \quad (26)$$

### 5.27 Rule ORI

Rule ORI is a rate rule for parameter ORI:

$$\frac{d}{dt}ORI = ks\_ori \cdot (Clb5 + epsilonori\_b2 \cdot Clb2) - kd\_ori \cdot ORI \quad (27)$$

### 5.28 Rule BUD

Rule BUD is a rate rule for parameter BUD:

$$\frac{d}{dt}BUD = ks\_bud \cdot (Cln2 + Cln3 + epsilonbud\_b5 \cdot Clb5) - kd\_bud \cdot BUD \quad (28)$$

### 5.29 Rule SPN

Rule SPN is a rate rule for parameter SPN:

$$\frac{d}{dt}SPN = \frac{ks\_spn \cdot Clb2}{J\_spn + Clb2} - kd\_spn \cdot SPN \quad (29)$$

## 6 Events

This is an overview of four events. Each event is initiated whenever its trigger condition switches from false to true. A delay function postpones the effects of an event to a later time point. At the time of execution, an event can assign values to species, parameters or compartments if these are not set to constant.

### 6.1 Event `Event_detection_for_END_M`

**Name** Event detection for END\_M

**Trigger condition** 
$$\text{SPN} \geq 1 \quad (30)$$

**Assignment** 
$$\text{END\_M} = \text{time} \quad (31)$$

### 6.2 Event `Event_detection_for_Cell_Division_and_BUD_SPN_reset`

**Name** Event detection for Cell Division and BUD/SPN reset

**Trigger condition** 
$$\text{Clb2} < 0.3 \quad (32)$$

**Assignments**

$$\text{mass} = \exp(1 \cdot \mu \cdot D) \cdot \text{mass} \quad (33)$$

$$\text{BUD} = 0 \quad (34)$$

$$\text{SPN} = 0 \quad (35)$$

### 6.3 Event `Event_detection_for_START_S`

**Name** Event detection for START\_S

**Trigger condition** 
$$\text{ORI} > 1 \quad (36)$$

**Assignments**

$$\text{START\_S} = \text{time} \quad (37)$$

$$\text{END\_M} = \text{time} + 1000 \quad (38)$$

### 6.4 Event `Event_detection_for_ORI_reset`

**Name** Event detection for ORI reset

**Trigger condition** 
$$\text{Clb2} + \text{Clb5} < 0.2 \quad (39)$$

**Assignment** 
$$\text{ORI} = 0 \quad (40)$$

SBML<sup>2</sup>TeX 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.

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