

## SBML Model Report

# Model name: “Mueller2015 - Hepatocyte proliferation, T160 phosphorylation of CDK2”



May 5, 2016

## 1 General Overview

This is a document in SBML Level 2 Version 4 format. This model was created by the following two authors: Audald Lloret i Villas<sup>1</sup> and Marcel Schilling<sup>2</sup> at March 19<sup>th</sup> 2015 at 4:40 p. m. and last time modified at March 20<sup>th</sup> 2015 at 3:41 p. m. Table 1 shows 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	2
species types	0	species	39
events	0	constraints	0
reactions	69	function definitions	69
global parameters	79	unit definitions	2
rules	30	initial assignments	0

## Model Notes

Mueller2015 - Hepatocyte proliferation, T160phosphorylation of CDK2

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This model is described in the article: [T160-phosphorylated CDK2 defines threshold for HGF-dependent proliferation in primary hepatocytes](#). Mueller S, Huard J, Waldow K, Huang X, D'Alessandro LA, Bohl S, Brner K, Grimm D, Klamt S, Klingmller U, Schilling M. Mol. Syst. Biol. 2015; 11(3): 795

Abstract:

Liver regeneration is a tightly controlled process mainly achieved by proliferation of usually quiescent hepatocytes. The specific molecular mechanisms ensuring cell division only in response to proliferative signals such as hepatocyte growth factor (HGF) are not fully understood. Here, we combined quantitative time-resolved analysis of primary mouse hepatocyte proliferation at the single cell and at the population level with mathematical modeling. We showed that numerous G1/S transition components are activated upon hepatocyte isolation whereas DNA replication only occurs upon additional HGF stimulation. In response to HGF, Cyclin:CDK complex formation was increased, p21 rather than p27 was regulated, and Rb expression was enhanced. Quantification of protein levels at the restriction point showed an excess of CDK2 over CDK4 and limiting amounts of the transcription factor E2F-1. Analysis with our mathematical model revealed that T160 phosphorylation of CDK2 correlated best with growth factor-dependent proliferation, which we validated experimentally on both the population and the single cell level. In conclusion, we identified CDK2 phosphorylation as a gate-keeping mechanism to maintain hepatocyte quiescence in the absence of HGF.

This model is hosted on [BioModels Database](#) and identified by: [BIOMD0000000568](#).

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## 2 Unit Definitions

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

### 2.1 Unit time

**Name** time

**Definition** 3600 s

### 2.2 Unit substance

**Name** substance

**Definition** nmol

### 2.3 Unit volume

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

**Definition** 1

### 2.4 Unit area

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

**Definition**  $\text{m}^2$

### 2.5 Unit length

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

**Definition** m

## 3 Compartments

This model contains two compartments.

Table 2: Properties of all compartments.

Id	Name	SBO	Spatial Dimensions	Size	Unit	Constant	Outside
cell	Cytoplasm		3	1	litre	<input checked="" type="checkbox"/>	
Nucleus	Nucleus		3	1	litre	<input checked="" type="checkbox"/>	

#### 3.1 Compartment cell

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

**Name** Cytoplasm

#### 3.2 Compartment Nucleus

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

**Name** Nucleus

## 4 Species

This model contains 39 species. The boundary condition of 15 of these species is set to `true` so that these species' amount cannot be changed by any reaction. Section 9 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 Condition
S4	@cyto::C2E(T160Ũ,b)	cell	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S10	@cyto::C4D1(b)	cell	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S12	@cyto::p21(b)	cell	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S19	@cyto::C4D1(b!1).p21(b!1)	cell	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S20	@cyto::C2E(T160Ũ,b!1).p21(b!1)	cell	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
hgf	HGF	cell	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input checked="" type="checkbox"/>
inhp53	inhp53	cell	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input checked="" type="checkbox"/>
inherk	inhERK	cell	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input checked="" type="checkbox"/>
inhakt	inhAKT	cell	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input checked="" type="checkbox"/>
inhc4d1	inhc4d1	cell	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input checked="" type="checkbox"/>
ObsTotCycECDK2- _obs	TotCycECDK2	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input checked="" type="checkbox"/>
ObsTotCDK2T160- _obs	TotCDK2T160	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input checked="" type="checkbox"/>
ObsTotCycDCDK4- _obs	TotCycDCDK4	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input checked="" type="checkbox"/>
ObsTotP21_obs	TotP21	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input checked="" type="checkbox"/>
ObsCDK2P21_obs	CDK2P21	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input checked="" type="checkbox"/>
ObsTotE2F_obs	TotE2F	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input checked="" type="checkbox"/>
ObsTotRb_obs	TotRb	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input checked="" type="checkbox"/>
ObsPhosRbS788_obs	PhosRbS788	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input checked="" type="checkbox"/>

Id	Name	Compartment	Derived Unit	Constant	Boundary Condition
ObsPhosRbS800_obs	PhosRbS800	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input checked="" type="checkbox"/>
ObsDNAContent_obs	DNAContent	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input checked="" type="checkbox"/>
S23	@nuc::C2E(T160P̃,b!1).p21(b!1)	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S18	@nuc::C2E(T160P̃,b)	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S3	@nuc::C2E(T160Ũ,b!1).p21(b!1)	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S13	@nuc::C2E(T160Ũ,b)	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S24	@nuc::C4D1(b!1).p21(b!1)	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S26	@nuc::C4D1(b)	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S5	@nuc::dnapre()	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S17	@nuc::dnapre1()	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S22	@nuc::dnapre2()	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S25	@nuc::dnapre3()	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S27	@nuc::dnapre4()	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S16	@nuc::e2f(b!1).rb(S788P̃,S800Ũ,b!1)	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S2	@nuc::e2f(b!1).rb(S788Ũ,S800Ũ,b!1)	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S14	@nuc::e2f(b)	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S11	@nuc::p21(b)	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S21	@nuc::rb(S788P̃,S800P̃,b)	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S15	@nuc::rb(S788P̃,S800Ũ,b)	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S1	@nuc::rb(S788Ũ,S800Ũ,b)	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>
S28	@nuc::dnapre5()	Nucleus	$\text{nmol} \cdot \text{l}^{-1}$	<input type="checkbox"/>	<input type="checkbox"/>

## 5 Parameters

This model contains 79 global parameters.

Table 4: Properties of each parameter.

Id	Name	SBO	Value	Unit	Constant
Vnuc	Vnuc		0.250		✓
Vcyto	Vcyto		12.670		✓
perk	perk		0.009		✓
nerk	nerk		1.147		✓
pakt	pakt		0.035		✓
nakt	nakt		1.096		✓
aerk	aerk		0.160		✓
aakt	aakt		0.530		✓
atf	atf		0.601		✓
ks_c4	ks_c4		14298.672		✓
kdeg_c4	kdeg_c4		1.014		✓
kdeg_c4gsk3b	kdeg_c4gsk3b		0.108		✓
ks_c2myc	ks_c2myc		0.158		✓
ks_c2e2f	ks_c2e2f		2.199		✓
kdeg-_c2c2gsk3b	kdeg_c2c2gsk3b		5.588		✓
kdeg_c2gsk3b	kdeg_c2gsk3b		$1.55090179808215 \cdot 10^{-5}$		✓
kdeg_c2	kdeg_c2		0.226		✓
kdp_c2cak	kdp_c2cak		101.282		✓
kc2cak	kc2cak		0.317		✓
ks_p21p53	ks_p21p53		$3.84136205729286 \cdot 10^{-6}$		✓
ks_p21e2f	ks_p21e2f		0.812		✓
kdeg-_p21erkskp2	kdeg_p21erkskp2		$2.82976267377082 \cdot 10^{-4}$		✓
kdeg-_p21c2skp2	kdeg_p21c2skp2		0.040		✓
kdeg_p21skp2	kdeg_p21skp2		0.751		✓
kdeg-_p21gsk3b	kdeg_p21gsk3b		0.005		✓
kdeg_p21erk	kdeg_p21erk		0.736		✓
Kd_p21c4	Kd_p21c4		99.997		✓
kb_p21c4	kb_p21c4		14.308		✓
Kd_p21c2	Kd_p21c2		0.010		✓
kb_p21c2	kb_p21c2		997.938		✓
ki	ki		0.092		✓
kinh_p21akt	kinh_p21akt		0.440		✓
ks_e2fe2f	ks_e2fe2f		0.460		✓

Id	Name	SBO	Value	Unit	Constant
ks_e2fmyc	ks_e2fmyc		$2.49174531457788 \cdot 10^{-6}$		<input checked="" type="checkbox"/>
kdege2fplus	kdege2fplus		$4.18153340918872 \cdot 10^{-5}$		<input checked="" type="checkbox"/>
kdeg- _e2fbound	kdeg_e2fbound		0.100		<input checked="" type="checkbox"/>
ks_rb	ks_rb		72.525		<input checked="" type="checkbox"/>
ks_rbe2f	ks_rbe2f		20.013		<input checked="" type="checkbox"/>
kdeg_rbp21	kdeg_rbp21		0.864		<input checked="" type="checkbox"/>
kdeg_rbbound	kdeg_rbbound		0.089		<input checked="" type="checkbox"/>
kdegrbplus	kdegrbplus		0.258		<input checked="" type="checkbox"/>
kb_rbe2f	kb_rbe2f		229.976		<input checked="" type="checkbox"/>
Kd_rb_e2f	Kd_rb_e2f		50.003		<input checked="" type="checkbox"/>
kb_rbpe2f	kb_rbpe2f		182.218		<input checked="" type="checkbox"/>
Kd_rbp_e2f	Kd_rbp_e2f		481.485		<input checked="" type="checkbox"/>
kcatprbc4	kcatprbc4		2797.823		<input checked="" type="checkbox"/>
kcatp_rbc2	kcatp_rbc2		7142308.072		<input checked="" type="checkbox"/>
kcatdp_rbc2	kcatdp_rbc2		0.003		<input checked="" type="checkbox"/>
kcatdp_rbc4	kcatdp_rbc4		2892.022		<input checked="" type="checkbox"/>
kinh_pp1	kinh_pp1		16634.940		<input checked="" type="checkbox"/>
Km_dprb	Km_dprb		0.119		<input checked="" type="checkbox"/>
Km_prb	Km_prb		2.035		<input checked="" type="checkbox"/>
nrb	nrb		3.000		<input checked="" type="checkbox"/>
k_dna	k_dna		0.009		<input checked="" type="checkbox"/>
k_delay	k_delay		23.666		<input checked="" type="checkbox"/>
Vratio	Vratio		0.020		<input type="checkbox"/>
erk	erk		0.160		<input type="checkbox"/>
akt	akt		0.530		<input type="checkbox"/>
gsk3b	gsk3b		0.470		<input type="checkbox"/>
tf	tf		0.635		<input type="checkbox"/>
tfp21	tfp21		0.635		<input type="checkbox"/>
kp_c2cak	kp_c2cak		101.599		<input type="checkbox"/>
kd_p21c4	kd_p21c4		1430.784		<input type="checkbox"/>
kd_p21c2	kd_p21c2		9.982		<input type="checkbox"/>
kimport	kimport		0.074		<input type="checkbox"/>
kdeg_e2ffree	kdeg_e2ffree		0.100		<input type="checkbox"/>
kdeg_rbfree	kdeg_rbfree		0.347		<input type="checkbox"/>
kd_rbe2f	kd_rbe2f		11499.401		<input type="checkbox"/>
kd_rbpe2f	kd_rbpe2f		87735.366		<input type="checkbox"/>
kcatp_rbc4	kcatp_rbc4		2797.823		<input type="checkbox"/>
scale- _TotcycDCDK4	scale-_TotcycDCDK4		0.565		<input checked="" type="checkbox"/>
scale- _TotcycECDK2	scale-_TotcycECDK2		0.189		<input checked="" type="checkbox"/>

Id	Name	SBO	Value	Unit	Constant
scale- _Totp21CDK2	scale_Totp21CDK2		0.340		✓
scale- _TotCDK2T160	scale- _TotCDK2T160		2.728		✓
scale_TotRb	scale_TotRb		0.261		✓
scale- _PhosRbS788	scale_PhosRbS788		0.674		✓
scale- _PhosRbS800	scale_PhosRbS800		0.824		✓
scale_Totp21	scale_Totp21		0.173		✓
scale_TotE2F	scale_TotE2F		28.742		✓

## 6 Function definitions

This is an overview of 69 function definitions.

### 6.1 Function definition [Function\\_for\\_reaction\\_1\\_0](#)

**Name** Function for reaction\_1

**Arguments**  $\text{vol}(\text{cell})$ ,  $\text{ks\_c4}$ ,  $\text{tf}$

**Mathematical Expression**

$$\frac{\text{ks\_c4} \cdot \text{tf}}{\text{vol}(\text{cell})} \quad (1)$$

### 6.2 Function definition [Function\\_for\\_reaction\\_2\\_0](#)

**Name** Function for reaction\_2

**Arguments**  $[\text{S14}]$ ,  $[\text{S16}]$ ,  $\text{vol}(\text{cell})$ ,  $\text{ks\_c2e2f}$ ,  $\text{ks\_c2myc}$ ,  $\text{tf}$

**Mathematical Expression**

$$\frac{\text{ks\_c2myc} \cdot \text{tf} + \text{ks\_c2e2f} \cdot ([\text{S14}] + [\text{S16}])}{\text{vol}(\text{cell})} \quad (2)$$

### 6.3 Function definition [Function\\_for\\_reaction\\_58\\_0](#)

**Name** Function for reaction\_58

**Arguments**  $[\text{S22}]$ ,  $\text{vol}(\text{cell})$ ,  $\text{k\_delay}$



**Mathematical Expression**

$$\frac{k\_delay \cdot [S22]}{vol(cell)} \quad (3)$$

**6.4 Function definition** [Function\\_for\\_reaction\\_59\\_0](#)**Name** Function for reaction\_59**Arguments** [S24], vol (cell), kdeg\_c4**Mathematical Expression**

$$\frac{kdeg\_c4 \cdot [S24]}{vol(cell)} \quad (4)$$

**6.5 Function definition** [Function\\_for\\_reaction\\_3\\_0](#)**Name** Function for reaction\_3**Arguments** [S3], vol (cell), gsk3b, kdeg\_c2, kdeg\_c2gsk3b**Mathematical Expression**

$$\frac{(kdeg\_c2 + kdeg\_c2gsk3b \cdot gsk3b) \cdot [S3]}{vol(cell)} \quad (5)$$

**6.6 Function definition** [Function\\_for\\_reaction\\_4\\_0](#)**Name** Function for reaction\_4**Arguments** [S4], vol (cell), gsk3b, kdeg\_c2, kdeg\_c2gsk3b**Mathematical Expression**

$$\frac{(kdeg\_c2 + kdeg\_c2gsk3b \cdot gsk3b) \cdot [S4]}{vol(cell)} \quad (6)$$

**6.7 Function definition** [Function\\_for\\_reaction\\_5\\_0](#)**Name** Function for reaction\_5**Arguments** [S14], vol (cell), ks\_p21e2f, ks\_p21p53, tfp21**Mathematical Expression**

$$\frac{(ks\_p21p53 + ks\_p21e2f \cdot [S14]) \cdot tfp21}{vol(cell)} \quad (7)$$

## 6.8 Function definition [Function\\_for\\_reaction\\_6\\_0](#)

**Name** Function for reaction\_6

**Arguments** [S3], vol (cell), kd\_p21c2

**Mathematical Expression**

$$\frac{\text{kd\_p21c2} \cdot [\text{S3}]}{\text{vol}(\text{cell})} \quad (8)$$

## 6.9 Function definition [Function\\_for\\_reaction\\_7\\_0](#)

**Name** Function for reaction\_7

**Arguments** [S14], [S18], [S3], vol (cell), erk, kdeg\_p21c2skp2, kdeg\_p21erkskp2, kdeg\_p21skp2

**Mathematical Expression**

$$\frac{(\text{kdeg\_p21erkskp2} \cdot \text{erk} + \text{kdeg\_p21c2skp2} \cdot [\text{S18}] + \text{kdeg\_p21skp2}) \cdot [\text{S14}] \cdot [\text{S3}]}{\text{vol}(\text{cell})} \quad (9)$$

## 6.10 Function definition [Function\\_for\\_reaction\\_8\\_0](#)

**Name** Function for reaction\_8

**Arguments** [S14], vol (cell), ks\_rb, ks\_rbe2f

**Mathematical Expression**

$$\frac{\text{ks\_rb} + \text{ks\_rbe2f} \cdot [\text{S14}]}{\text{vol}(\text{cell})} \quad (10)$$

## 6.11 Function definition [Function\\_for\\_reaction\\_9\\_0](#)

**Name** Function for reaction\_9

**Arguments** [S1], vol (cell), kdeg\_rbfree

**Mathematical Expression**

$$\frac{\text{kdeg\_rbfree} \cdot [\text{S1}]}{\text{vol}(\text{cell})} \quad (11)$$

## 6.12 Function definition [Function\\_for\\_reaction\\_10\\_0](#)

**Name** Function for reaction\_10

**Arguments** [S2], vol (cell), kdeg\_rbbound

**Mathematical Expression**

$$\frac{\text{kdeg\_rbbound} \cdot [\text{S2}]}{\text{vol}(\text{cell})} \quad (12)$$

### 6.13 Function definition [Function\\_for\\_reaction\\_11\\_0](#)

**Name** Function for reaction\_11

**Arguments** [S1], [S11], vol(cell), kdeg\_rbp21

**Mathematical Expression**

$$\frac{kdeg\_rbp21 \cdot [S11] \cdot [S1]}{vol(cell)} \quad (13)$$

### 6.14 Function definition [Function\\_for\\_reaction\\_12\\_0](#)

**Name** Function for reaction\_12

**Arguments** [S11], [S2], vol(cell), kdeg\_rbp21

**Mathematical Expression**

$$\frac{kdeg\_rbp21 \cdot [S11] \cdot [S2]}{vol(cell)} \quad (14)$$

### 6.15 Function definition [Function\\_for\\_reaction\\_13\\_0](#)

**Name** Function for reaction\_13

**Arguments** [S14], vol(cell), ks\_e2fe2f, ks\_e2fmyc, tf

**Mathematical Expression**

$$\frac{(ks\_e2fe2f \cdot [S14] + ks\_e2fmyc) \cdot tf}{vol(cell)} \quad (15)$$

### 6.16 Function definition [Function\\_for\\_reaction\\_14\\_0](#)

**Name** Function for reaction\_14

**Arguments** [S2], vol(cell), kdeg\_e2fbound

**Mathematical Expression**

$$\frac{kdeg\_e2fbound \cdot [S2]}{vol(cell)} \quad (16)$$

### 6.17 Function definition [Function\\_for\\_reaction\\_15\\_0](#)

**Name** Function for reaction\_15

**Arguments** [S2], vol(cell), kd\_rbe2f

**Mathematical Expression**

$$\frac{kd\_rbe2f \cdot [S2]}{vol(cell)} \quad (17)$$

### 6.18 Function definition [Function\\_for\\_reaction\\_16\\_0](#)

**Name** Function for reaction\_16

**Arguments** Km\_prb, [S1], [S24], vol (cell), kcatp\_rbc4, nrb

**Mathematical Expression**

$$\frac{\frac{kcatp\_rbc4 \cdot [S24] \cdot [S1]^{nrb}}{Km\_prb^{nrb} + [S1]^{nrb}}}{vol (cell)} \quad (18)$$

### 6.19 Function definition [Function\\_for\\_reaction\\_17\\_0](#)

**Name** Function for reaction\_17

**Arguments** Km\_prb, [S2], [S24], vol (cell), kcatp\_rbc4, nrb

**Mathematical Expression**

$$\frac{\frac{kcatp\_rbc4 \cdot [S24] \cdot [S2]^{nrb}}{Km\_prb^{nrb} + [S2]^{nrb}}}{vol (cell)} \quad (19)$$

### 6.20 Function definition [Function\\_for\\_reaction\\_18\\_0](#)

**Name** Function for reaction\_18

**Arguments** [S14], [S18], [S5], vol (cell), k\_dna

**Mathematical Expression**

$$\frac{k\_dna \cdot [S18] \cdot [S14] \cdot [S5]}{vol (cell)} \quad (20)$$

### 6.21 Function definition [Function\\_for\\_reaction\\_19\\_0](#)

**Name** Function for reaction\_19

**Arguments** [S10], vol (cell), gsk3b, kdeg\_c4, kdeg\_c4gsk3b

**Mathematical Expression**

$$\frac{(kdeg\_c4 + kdeg\_c4gsk3b \cdot gsk3b) \cdot [S10]}{vol (cell)} \quad (21)$$

## 6.22 Function definition [Function\\_for\\_reaction\\_20\\_0](#)

**Name** Function for reaction\_20

**Arguments** [S13], vol(cell), gsk3b, kdeg\_c2, kdeg\_c2gsk3b

**Mathematical Expression**

$$\frac{(kdeg\_c2 + kdeg\_c2gsk3b \cdot gsk3b) \cdot [S13]}{vol(cell)} \quad (22)$$

## 6.23 Function definition [Function\\_for\\_reaction\\_21\\_0](#)

**Name** Function for reaction\_21

**Arguments** [S13], vol(cell), kp\_c2cak

**Mathematical Expression**

$$\frac{kp\_c2cak \cdot [S13]}{vol(cell)} \quad (23)$$

## 6.24 Function definition [Function\\_for\\_reaction\\_22\\_0](#)

**Name** Function for reaction\_22

**Arguments** [S10], [S12], vol(cell), kb\_p21c4

**Mathematical Expression**

$$\frac{kb\_p21c4 \cdot [S10] \cdot [S12]}{vol(cell)} \quad (24)$$

## 6.25 Function definition [Function\\_for\\_reaction\\_23\\_0](#)

**Name** Function for reaction\_23

**Arguments** [S12], [S4], vol(cell), kb\_p21c2

**Mathematical Expression**

$$\frac{kb\_p21c2 \cdot [S4] \cdot [S12]}{vol(cell)} \quad (25)$$

## 6.26 Function definition [Function\\_for\\_reaction\\_24\\_0](#)

**Name** Function for reaction\_24

**Arguments** [S11], [S13], vol(cell), kb\_p21c2

**Mathematical Expression**

$$\frac{kb\_p21c2 \cdot [S11] \cdot [S13]}{vol(cell)} \quad (26)$$

### 6.27 Function definition [Function\\_for\\_reaction\\_25\\_0](#)

**Name** Function for reaction\_25

**Arguments** [S12], Vratio, vol (cell), kimport

**Mathematical Expression**

$$\frac{\frac{kimport}{Vratio} \cdot [S12]}{vol (cell)} \quad (27)$$

### 6.28 Function definition [Function\\_for\\_reaction\\_26\\_0](#)

**Name** Function for reaction\_26

**Arguments** [S12], Vratio, vol (cell), kimport

**Mathematical Expression**

$$\frac{kimport \cdot \left(1 - \frac{1}{Vratio}\right) \cdot [S12]}{vol (cell)} \quad (28)$$

### 6.29 Function definition [Function\\_for\\_reaction\\_27\\_0](#)

**Name** Function for reaction\_27

**Arguments** [S11], vol (cell), erk, gsk3b, kdeg\_p21erk, kdeg\_p21gsk3b

**Mathematical Expression**

$$\frac{(kdeg\_p21gsk3b \cdot gsk3b + kdeg\_p21erk \cdot erk) \cdot [S11]}{vol (cell)} \quad (29)$$

### 6.30 Function definition [Function\\_for\\_reaction\\_28\\_0](#)

**Name** Function for reaction\_28

**Arguments** [S12], vol (cell), erk, gsk3b, kdeg\_p21erk, kdeg\_p21gsk3b

**Mathematical Expression**

$$\frac{(kdeg\_p21gsk3b \cdot gsk3b + kdeg\_p21erk \cdot erk) \cdot [S12]}{vol (cell)} \quad (30)$$

### 6.31 Function definition [Function\\_for\\_reaction\\_29\\_0](#)

**Name** Function for reaction\_29

**Arguments** [S15], vol(cell), kdeg\_rbfree

**Mathematical Expression**

$$\frac{\text{kdeg\_rbfree} \cdot [\text{S15}]}{\text{vol}(\text{cell})} \quad (31)$$

### 6.32 Function definition [Function\\_for\\_reaction\\_30\\_0](#)

**Name** Function for reaction\_30

**Arguments** [S16], vol(cell), kdeg\_rbbound

**Mathematical Expression**

$$\frac{\text{kdeg\_rbbound} \cdot [\text{S16}]}{\text{vol}(\text{cell})} \quad (32)$$

### 6.33 Function definition [Function\\_for\\_reaction\\_31\\_0](#)

**Name** Function for reaction\_31

**Arguments** [S11], [S15], vol(cell), kdeg\_rbp21

**Mathematical Expression**

$$\frac{\text{kdeg\_rbp21} \cdot [\text{S11}] \cdot [\text{S15}]}{\text{vol}(\text{cell})} \quad (33)$$

### 6.34 Function definition [Function\\_for\\_reaction\\_32\\_0](#)

**Name** Function for reaction\_32

**Arguments** [S11], [S16], vol(cell), kdeg\_rbp21

**Mathematical Expression**

$$\frac{\text{kdeg\_rbp21} \cdot [\text{S11}] \cdot [\text{S16}]}{\text{vol}(\text{cell})} \quad (34)$$

### 6.35 Function definition [Function\\_for\\_reaction\\_33\\_0](#)

**Name** Function for reaction\_33

**Arguments** [S14], vol(cell), kdeg\_e2ffree

**Mathematical Expression**

$$\frac{\text{kdeg\_e2ffree} \cdot [\text{S14}]}{\text{vol}(\text{cell})} \quad (35)$$

### 6.36 Function definition [Function\\_for\\_reaction\\_34\\_0](#)

**Name** Function for reaction\_34

**Arguments** [S16], vol (cell), kdeg\_e2fbound

**Mathematical Expression**

$$\frac{kdeg\_e2fbound \cdot [S16]}{vol (cell)} \quad (36)$$

### 6.37 Function definition [Function\\_for\\_reaction\\_35\\_0](#)

**Name** Function for reaction\_35

**Arguments** [S1], [S14], vol (cell), kb\_rbe2f

**Mathematical Expression**

$$\frac{kb\_rbe2f \cdot [S1] \cdot [S14]}{vol (cell)} \quad (37)$$

### 6.38 Function definition [Function\\_for\\_reaction\\_36\\_0](#)

**Name** Function for reaction\_36

**Arguments** [S14], [S15], vol (cell), kb\_rbe2f

**Mathematical Expression**

$$\frac{kb\_rbe2f \cdot [S14] \cdot [S15]}{vol (cell)} \quad (38)$$

### 6.39 Function definition [Function\\_for\\_reaction\\_37\\_0](#)

**Name** Function for reaction\_37

**Arguments** [S16], vol (cell), kd\_rbe2f

**Mathematical Expression**

$$\frac{kd\_rbe2f \cdot [S16]}{vol (cell)} \quad (39)$$

### 6.40 Function definition [Function\\_for\\_reaction\\_38\\_0](#)

**Name** Function for reaction\_38

**Arguments** Km\_prb, [S15], [S18], vol (cell), kcatp\_rbc2, nrb

**Mathematical Expression**

$$\frac{kcatp\_rbc2 \cdot [S18] \cdot [S15]^{nrb}}{Km\_prb^{nrb} + [S15]^{nrb}} \cdot vol (cell) \quad (40)$$



#### 6.41 Function definition [Function\\_for\\_reaction\\_39\\_0](#)

**Name** Function for reaction\_39

**Arguments** Km\_prb, [S16], [S18], vol (cell), kcatp\_rbc2, nrb

**Mathematical Expression**

$$\frac{\frac{kcatp\_rbc2 \cdot [S18] \cdot [S16]^{nrb}}{Km\_prb^{nrb} + [S16]^{nrb}}}{vol (cell)} \quad (41)$$

#### 6.42 Function definition [Function\\_for\\_reaction\\_40\\_0](#)

**Name** Function for reaction\_40

**Arguments** Km\_dprb, [S15], [S18], vol (cell), kcatdp\_rbc4, kinh\_pp1, nrb

**Mathematical Expression**

$$\frac{\frac{\frac{kcatdp\_rbc4 \cdot [S15]^{nrb}}{Km\_dprb^{nrb} + [S15]^{nrb}} \cdot 1}{1 + kinh\_pp1 \cdot [S18]}}{vol (cell)} \quad (42)$$

#### 6.43 Function definition [Function\\_for\\_reaction\\_41\\_0](#)

**Name** Function for reaction\_41

**Arguments** Km\_dprb, [S16], [S18], vol (cell), kcatdp\_rbc4, kinh\_pp1, nrb

**Mathematical Expression**

$$\frac{\frac{\frac{kcatdp\_rbc4 \cdot [S16]^{nrb}}{Km\_dprb^{nrb} + [S16]^{nrb}} \cdot 1}{1 + kinh\_pp1 \cdot [S18]}}{vol (cell)} \quad (43)$$

#### 6.44 Function definition [Function\\_for\\_reaction\\_42\\_0](#)

**Name** Function for reaction\_42

**Arguments** [S17], vol (cell), k\_delay

**Mathematical Expression**

$$\frac{k\_delay \cdot [S17]}{vol (cell)} \quad (44)$$

#### 6.45 Function definition [Function\\_for\\_reaction\\_43\\_0](#)

**Name** Function for reaction\_43

**Arguments** [S19], vol (cell), gsk3b, kdeg\_c4, kdeg\_c4gsk3b

**Mathematical Expression**

$$\frac{(kdeg\_c4 + kdeg\_c4gsk3b \cdot gsk3b) \cdot [S19]}{vol (cell)} \quad (45)$$

#### 6.46 Function definition [Function\\_for\\_reaction\\_44\\_0](#)

**Name** Function for reaction\_44

**Arguments** [S18], vol (cell), gsk3b, kdeg\_c2, kdeg\_c2gsk3b

**Mathematical Expression**

$$\frac{(kdeg\_c2 + kdeg\_c2gsk3b \cdot gsk3b) \cdot [S18]}{vol (cell)} \quad (46)$$

#### 6.47 Function definition [Function\\_for\\_reaction\\_45\\_0](#)

**Name** Function for reaction\_45

**Arguments** [S20], vol (cell), gsk3b, kdeg\_c2, kdeg\_c2gsk3b

**Mathematical Expression**

$$\frac{(kdeg\_c2 + kdeg\_c2gsk3b \cdot gsk3b) \cdot [S20]}{vol (cell)} \quad (47)$$

#### 6.48 Function definition [Function\\_for\\_reaction\\_46\\_0](#)

**Name** Function for reaction\_46

**Arguments** [S18], vol (cell), gsk3b, kdeg\_c2c2gsk3b

**Mathematical Expression**

$$\frac{kdeg\_c2c2gsk3b \cdot gsk3b \cdot [S18]}{vol (cell)} \quad (48)$$

#### 6.49 Function definition [Function\\_for\\_reaction\\_47\\_0](#)

**Name** Function for reaction\_47

**Arguments** [S18], vol (cell), kdp\_c2cak

**Mathematical Expression**

$$\frac{\text{kdp\_c2cak} \cdot [\text{S18}]}{\text{vol}(\text{cell})} \quad (49)$$

#### 6.50 Function definition [Function\\_for\\_reaction\\_48\\_0](#)

**Name** Function for reaction\_48

**Arguments** [S19], vol (cell), kd\_p21c4

**Mathematical Expression**

$$\frac{\text{kd\_p21c4} \cdot [\text{S19}]}{\text{vol}(\text{cell})} \quad (50)$$

#### 6.51 Function definition [Function\\_for\\_reaction\\_49\\_0](#)

**Name** Function for reaction\_49

**Arguments** [S11], [S18], vol (cell), kb\_p21c2

**Mathematical Expression**

$$\frac{\text{kb\_p21c2} \cdot [\text{S11}] \cdot [\text{S18}]}{\text{vol}(\text{cell})} \quad (51)$$

#### 6.52 Function definition [Function\\_for\\_reaction\\_50\\_0](#)

**Name** Function for reaction\_50

**Arguments** [S20], vol (cell), kd\_p21c2

**Mathematical Expression**

$$\frac{\text{kd\_p21c2} \cdot [\text{S20}]}{\text{vol}(\text{cell})} \quad (52)$$

#### 6.53 Function definition [Function\\_for\\_reaction\\_51\\_0](#)

**Name** Function for reaction\_51

**Arguments** [S19], Vratio, vol (cell), kimport

**Mathematical Expression**

$$\frac{\frac{\text{kimport}}{\text{Vratio}} \cdot [\text{S19}]}{\text{vol}(\text{cell})} \quad (53)$$

#### 6.54 Function definition [Function\\_for\\_reaction\\_52\\_0](#)

**Name** Function for reaction\_52

**Arguments** [S20], Vratio, vol (cell), kimport

**Mathematical Expression**

$$\frac{\frac{kimport}{Vratio} \cdot [S20]}{vol (cell)} \quad (54)$$

#### 6.55 Function definition [Function\\_for\\_reaction\\_53\\_0](#)

**Name** Function for reaction\_53

**Arguments** [S19], Vratio, vol (cell), kimport

**Mathematical Expression**

$$\frac{kimport \cdot \left(1 - \frac{1}{Vratio}\right) \cdot [S19]}{vol (cell)} \quad (55)$$

#### 6.56 Function definition [Function\\_for\\_reaction\\_54\\_0](#)

**Name** Function for reaction\_54

**Arguments** [S20], Vratio, vol (cell), kimport

**Mathematical Expression**

$$\frac{kimport \cdot \left(1 - \frac{1}{Vratio}\right) \cdot [S20]}{vol (cell)} \quad (56)$$

#### 6.57 Function definition [Function\\_for\\_reaction\\_55\\_0](#)

**Name** Function for reaction\_55

**Arguments** [S21], vol (cell), kdeg\_rbfree

**Mathematical Expression**

$$\frac{kdeg\_rbfree \cdot [S21]}{vol (cell)} \quad (57)$$

### 6.58 Function definition [Function\\_for\\_reaction\\_56\\_0](#)

**Name** Function for reaction\_56

**Arguments** [S11], [S21], vol (cell), kdeg\_rbp21

**Mathematical Expression**

$$\frac{kdeg\_rbp21 \cdot [S11] \cdot [S21]}{vol (cell)} \quad (58)$$

### 6.59 Function definition [Function\\_for\\_reaction\\_57\\_0](#)

**Name** Function for reaction\_57

**Arguments** Km\_dprb, [S18], [S21], vol (cell), kcatdp\_rbc2, kinh\_pp1, nrb

**Mathematical Expression**

$$\frac{\frac{kcatdp\_rbc2 \cdot [S21]^{nrb}}{Km\_dprb^{nrb} + [S21]^{nrb}} \cdot 1}{1 + kinh\_pp1 \cdot [S18]} \cdot vol (cell) \quad (59)$$

### 6.60 Function definition [Function\\_for\\_reaction\\_60\\_0](#)

**Name** Function for reaction\_60

**Arguments** [S23], vol (cell), gsk3b, kdeg\_c2, kdeg\_c2gsk3b

**Mathematical Expression**

$$\frac{(kdeg\_c2 + kdeg\_c2gsk3b \cdot gsk3b) \cdot [S23]}{vol (cell)} \quad (60)$$

### 6.61 Function definition [Function\\_for\\_reaction\\_61\\_0](#)

**Name** Function for reaction\_61

**Arguments** [S24], vol (cell), kd\_p21c4

**Mathematical Expression**

$$\frac{kd\_p21c4 \cdot [S24]}{vol (cell)} \quad (61)$$

## 6.62 Function definition [Function\\_for\\_reaction\\_62\\_0](#)

**Name** Function for reaction\_62

**Arguments** [S23], vol (cell), kd\_p21c2

**Mathematical Expression**

$$\frac{\text{kd\_p21c2} \cdot [\text{S23}]}{\text{vol}(\text{cell})} \quad (62)$$

## 6.63 Function definition [Function\\_for\\_reaction\\_63\\_0](#)

**Name** Function for reaction\_63

**Arguments** [S14], [S18], [S23], vol (cell), erk, kdeg\_p21c2skp2, kdeg\_p21erkskp2, kdeg\_p21skp2

**Mathematical Expression**

$$\frac{(\text{kdeg\_p21erkskp2} \cdot \text{erk} + \text{kdeg\_p21c2skp2} \cdot [\text{S18}] + \text{kdeg\_p21skp2}) \cdot [\text{S14}] \cdot [\text{S23}]}{\text{vol}(\text{cell})} \quad (63)$$

## 6.64 Function definition [Function\\_for\\_reaction\\_64\\_0](#)

**Name** Function for reaction\_64

**Arguments** [S14], [S18], [S24], vol (cell), erk, kdeg\_p21c2skp2, kdeg\_p21erkskp2, kdeg\_p21skp2

**Mathematical Expression**

$$\frac{(\text{kdeg\_p21erkskp2} \cdot \text{erk} + \text{kdeg\_p21c2skp2} \cdot [\text{S18}] + \text{kdeg\_p21skp2}) \cdot [\text{S14}] \cdot [\text{S24}]}{\text{vol}(\text{cell})} \quad (64)$$

## 6.65 Function definition [Function\\_for\\_reaction\\_65\\_0](#)

**Name** Function for reaction\_65

**Arguments** [S25], vol (cell), k\_delay

**Mathematical Expression**

$$\frac{\text{k\_delay} \cdot [\text{S25}]}{\text{vol}(\text{cell})} \quad (65)$$

### 6.66 Function definition [Function\\_for\\_reaction\\_66\\_0](#)

**Name** Function for reaction\_66

**Arguments** [S26], vol (cell), gsk3b, kdeg\_c4, kdeg\_c4gsk3b

**Mathematical Expression**

$$\frac{(kdeg\_c4 + kdeg\_c4gsk3b \cdot gsk3b) \cdot [S26]}{vol (cell)} \quad (66)$$

### 6.67 Function definition [Function\\_for\\_reaction\\_67\\_0](#)

**Name** Function for reaction\_67

**Arguments** [S11], [S26], vol (cell), kb\_p21c4

**Mathematical Expression**

$$\frac{kb\_p21c4 \cdot [S11] \cdot [S26]}{vol (cell)} \quad (67)$$

### 6.68 Function definition [Function\\_for\\_reaction\\_68\\_0](#)

**Name** Function for reaction\_68

**Arguments** [S27], vol (cell), k\_delay

**Mathematical Expression**

$$\frac{k\_delay \cdot [S27]}{vol (cell)} \quad (68)$$

### 6.69 Function definition [Function\\_for\\_reaction\\_69\\_0](#)

**Name** Function for reaction\_69

**Arguments** [S28], vol (cell), k\_delay

**Mathematical Expression**

$$\frac{k\_delay \cdot [S28]}{vol (cell)} \quad (69)$$

## 7 Rules

This is an overview of 30 rules.

### 7.1 Rule ObsCDK2P21\_obs

Rule ObsCDK2P21\_obs is an assignment rule for species ObsCDK2P21\_obs:

$$\text{ObsCDK2P21\_obs} = \frac{\text{scale\_Totp21CDK2} \cdot (\text{Vnuc} \cdot ([\text{S3}] + [\text{S23}]) + \text{Vcyto} \cdot [\text{S20}])}{\text{Vnuc} + \text{Vcyto}} \quad (70)$$

### 7.2 Rule ObsTotE2F\_obs

Rule ObsTotE2F\_obs is an assignment rule for species ObsTotE2F\_obs:

$$\text{ObsTotE2F\_obs} = \frac{(\text{scale\_TotE2F} + \text{scale\_TotRb}) \cdot \text{Vnuc} \cdot ([\text{S2}] + [\text{S14}] + [\text{S16}])}{\text{Vnuc} + \text{Vcyto}} \quad (71)$$

### 7.3 Rule hgf

Rule hgf is an assignment rule for species hgf:

$$\text{hgf} = \begin{cases} 0 & \text{if time} < 1 \\ \begin{cases} 0 & \text{if time} < 24 \\ 1 & \text{otherwise} \end{cases} & \text{otherwise} \end{cases} \quad (72)$$

### 7.4 Rule inhp53

Rule inhp53 is an assignment rule for species inhp53:

$$\text{inhp53} = \begin{cases} 0 & \text{if time} < 1 \\ \begin{cases} 0 & \text{if time} < 0 \\ 0 & \text{otherwise} \end{cases} & \text{otherwise} \end{cases} \quad (73)$$

### 7.5 Rule inherk

Rule inherk is an assignment rule for species inherk:

$$\text{inherk} = \begin{cases} 0 & \text{if time} < 1 \\ \begin{cases} 0 & \text{if time} < 0 \\ 0 & \text{otherwise} \end{cases} & \text{otherwise} \end{cases} \quad (74)$$

### 7.6 Rule inhakt

Rule inhakt is an assignment rule for species inhakt:

$$\text{inhakt} = \begin{cases} 0 & \text{if time} < 1 \\ \begin{cases} 0 & \text{if time} < 0 \\ 0 & \text{otherwise} \end{cases} & \text{otherwise} \end{cases} \quad (75)$$



### 7.7 Rule `inhc4d1`

Rule `inhc4d1` is an assignment rule for species `inhc4d1`:

$$\text{inhc4d1} = \begin{cases} 0 & \text{if time} < 1 \\ \begin{cases} 0 & \text{if time} < 0 \\ 0 & \text{otherwise} \end{cases} & \text{otherwise} \end{cases} \quad (76)$$

### 7.8 Rule `ObsTotCycECDK2_obs`

Rule `ObsTotCycECDK2_obs` is an assignment rule for species `ObsTotCycECDK2_obs`:

$$\begin{aligned} \text{ObsTotCycECDK2\_obs} \\ = \frac{\text{scale\_TotcycECDK2} \cdot (\text{Vnuc} \cdot ([S3] + [S13] + [S18] + [S23]) + \text{Vcyto} \cdot ([S4] + [S20]))}{\text{Vnuc} + \text{Vcyto}} \end{aligned} \quad (77)$$

### 7.9 Rule `ObsTotCDK2T160_obs`

Rule `ObsTotCDK2T160_obs` is an assignment rule for species `ObsTotCDK2T160_obs`:

$$\text{ObsTotCDK2T160\_obs} = \frac{\text{scale\_TotCDK2T160} \cdot \text{Vnuc} \cdot ([S18] + [S23])}{\text{Vnuc} + \text{Vcyto}} \quad (78)$$

### 7.10 Rule `ObsTotCycDCDK4_obs`

Rule `ObsTotCycDCDK4_obs` is an assignment rule for species `ObsTotCycDCDK4_obs`:

$$\text{ObsTotCycDCDK4\_obs} = \frac{\text{scale\_TotcycDCDK4} \cdot (\text{Vnuc} \cdot [S24] + \text{Vcyto} \cdot [S19])}{\text{Vnuc} + \text{Vcyto}} \quad (79)$$

### 7.11 Rule `ObsTotP21_obs`

Rule `ObsTotP21_obs` is an assignment rule for species `ObsTotP21_obs`:

$$\begin{aligned} \text{ObsTotP21\_obs} \\ = \frac{\text{scale\_Totp21} \cdot (\text{Vnuc} \cdot ([S3] + [S11] + [S23] + [S24]) + \text{Vcyto} \cdot ([S12] + [S19] + [S20]))}{\text{Vnuc} + \text{Vcyto}} \end{aligned} \quad (80)$$

### 7.12 Rule `ObsTotRb_obs`

Rule `ObsTotRb_obs` is an assignment rule for species `ObsTotRb_obs`:

$$\text{ObsTotRb\_obs} = \frac{\text{scale\_TotRb} \cdot \text{Vnuc} \cdot ([S1] + [S2] + [S15] + [S16] + [S21])}{\text{Vnuc} + \text{Vcyto}} \quad (81)$$

### 7.13 Rule `ObsPhosRbS788_obs`

Rule `ObsPhosRbS788_obs` is an assignment rule for species `ObsPhosRbS788_obs`:

$$\text{ObsPhosRbS788\_obs} = \frac{\text{scale\_PhosRbS788} \cdot \text{Vnuc} \cdot ([\text{S15}] + [\text{S16}] + [\text{S21}])}{\text{Vnuc} + \text{Vcyto}} \quad (82)$$

### 7.14 Rule `ObsPhosRbS800_obs`

Rule `ObsPhosRbS800_obs` is an assignment rule for species `ObsPhosRbS800_obs`:

$$\text{ObsPhosRbS800\_obs} = \frac{\text{scale\_PhosRbS800} \cdot \text{Vnuc} \cdot [\text{S21}]}{\text{Vnuc} + \text{Vcyto}} \quad (83)$$

### 7.15 Rule `ObsDNAContent_obs`

Rule `ObsDNAContent_obs` is an assignment rule for species `ObsDNAContent_obs`:

$$\text{ObsDNAContent\_obs} = 2 - ([\text{S5}] + [\text{S17}] + [\text{S22}] + [\text{S25}] + [\text{S27}] + [\text{S28}]) \quad (84)$$

### 7.16 Rule `Vratio`

Rule `Vratio` is an assignment rule for parameter `Vratio`:

$$\text{Vratio} = \frac{\text{Vnuc}}{\text{Vcyto}} \quad (85)$$

### 7.17 Rule `erk`

Rule `erk` is an assignment rule for parameter `erk`:

$$\text{erk} = (1 - [\text{inherk}]) \cdot \left( \frac{(1 - \text{aerk}) \cdot (\text{perk}^{\text{nerk}} + 1) \cdot [\text{hgf}]^{\text{nerk}}}{[\text{hgf}]^{\text{nerk}} + \text{perk}^{\text{nerk}}} + \text{aerk} \right) \quad (86)$$

### 7.18 Rule `akt`

Rule `akt` is an assignment rule for parameter `akt`:

$$\text{akt} = (1 - [\text{inhakt}]) \cdot \left( \frac{(1 - \text{aakt}) \cdot (\text{pakt}^{\text{nakt}} + 1) \cdot [\text{hgf}]^{\text{nakt}}}{[\text{hgf}]^{\text{nakt}} + \text{pakt}^{\text{nakt}}} + \text{aakt} \right) \quad (87)$$

### 7.19 Rule `gsk3b`

Rule `gsk3b` is an assignment rule for parameter `gsk3b`:

$$\text{gsk3b} = 1 - \text{akt} \quad (88)$$

### 7.20 Rule `tf`

Rule `tf` is an assignment rule for parameter `tf`:

$$tf = (1 - atf) \cdot erk \cdot (1 - gsk3b) + atf \quad (89)$$

### 7.21 Rule `tfp21`

Rule `tfp21` is an assignment rule for parameter `tfp21`:

$$tfp21 = (1 - [inhp53]) \cdot tf \quad (90)$$

### 7.22 Rule `kp_c2cak`

Rule `kp_c2cak` is an assignment rule for parameter `kp_c2cak`:

$$kp\_c2cak = kdp\_c2cak + kc2cak \quad (91)$$

### 7.23 Rule `kd_p21c4`

Rule `kd_p21c4` is an assignment rule for parameter `kd_p21c4`:

$$kd\_p21c4 = Kd\_p21c4 \cdot kb\_p21c4 \quad (92)$$

### 7.24 Rule `kd_p21c2`

Rule `kd_p21c2` is an assignment rule for parameter `kd_p21c2`:

$$kd\_p21c2 = Kd\_p21c2 \cdot kb\_p21c2 \quad (93)$$

### 7.25 Rule `kimport`

Rule `kimport` is an assignment rule for parameter `kimport`:

$$kimport = \frac{ki}{1 + kinh\_p21akt \cdot akt} \quad (94)$$

### 7.26 Rule `kdeg_e2ffree`

Rule `kdeg_e2ffree` is an assignment rule for parameter `kdeg_e2ffree`:

$$kdeg\_e2ffree = kdeg\_e2fbound + kdege2fplus \quad (95)$$

### 7.27 Rule `kdeg_rbfree`

Rule `kdeg_rbfree` is an assignment rule for parameter `kdeg_rbfree`:

$$kdeg\_rbfree = kdeg\_rbbound + kdeg_rbplus \quad (96)$$

### 7.28 Rule `kd_rbe2f`

Rule `kd_rbe2f` is an assignment rule for parameter `kd_rbe2f`:

$$\text{kd\_rbe2f} = \text{kb\_rbe2f} \cdot \text{Kd\_rb\_e2f} \quad (97)$$

### 7.29 Rule `kd_rbpe2f`

Rule `kd_rbpe2f` is an assignment rule for parameter `kd_rbpe2f`:

$$\text{kd\_rbpe2f} = \text{kb\_rbpe2f} \cdot \text{Kd\_rbp\_e2f} \quad (98)$$

### 7.30 Rule `kcatp_rbc4`

Rule `kcatp_rbc4` is an assignment rule for parameter `kcatp_rbc4`:

$$\text{kcatp\_rbc4} = \text{kcatprbc4} \cdot (1 - [\text{inhc4d1}]) \quad (99)$$

## 8 Reactions

This model contains 69 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 5: Overview of all reactions

Nº	Id	Name	Reaction Equation	SBO
1	reaction_1	reaction_1	$\emptyset \longrightarrow S10$	
2	reaction_2	reaction_2	$\emptyset \xrightarrow{S14, S16} S4$	
3	reaction_3	reaction_3	$S3 \xrightarrow{S3} S11$	
4	reaction_4	reaction_4	$S4 \xrightarrow{S4} \emptyset$	
5	reaction_5	reaction_5	$\emptyset \xrightarrow{S14} S12$	
6	reaction_6	reaction_6	$S3 \xrightarrow{S3} S11 + S13$	
7	reaction_7	reaction_7	$S3 \xrightarrow{S18, S14, S3} S13$	
8	reaction_8	reaction_8	$\emptyset \xrightarrow{S14} S1$	
9	reaction_9	reaction_9	$S1 \xrightarrow{S1} \emptyset$	
10	reaction_10	reaction_10	$S2 \xrightarrow{S2} S14$	
11	reaction_11	reaction_11	$S1 \xrightarrow{S11, S1} \emptyset$	
12	reaction_12	reaction_12	$S2 \xrightarrow{S11, S2} S14$	
13	reaction_13	reaction_13	$\emptyset \xrightarrow{S14} S14$	
14	reaction_14	reaction_14	$S2 \xrightarrow{S2} S1$	
15	reaction_15	reaction_15	$S2 \xrightarrow{S2} S1 + S14$	
16	reaction_16	reaction_16	$S1 \xrightarrow{S24, S1} S15$	
17	reaction_17	reaction_17	$S2 \xrightarrow{S24, S2} S16$	

Nº	Id	Name	Reaction Equation	SBO
18	reaction_18	reaction_18	$S5 \xrightarrow{S18, S14, S5} S17$	
19	reaction_19	reaction_19	$S10 \xrightarrow{S10} \emptyset$	
20	reaction_20	reaction_20	$S13 \xrightarrow{S13} \emptyset$	
21	reaction_21	reaction_21	$S13 \xrightarrow{S13} S18$	
22	reaction_22	reaction_22	$S10 + S12 \xrightarrow{S10, S12} S19$	
23	reaction_23	reaction_23	$S4 + S12 \xrightarrow{S4, S12} S20$	
24	reaction_24	reaction_24	$S11 + S13 \xrightarrow{S11, S13} S3$	
25	reaction_25	reaction_25	$S12 \xrightarrow{S12} S11$	
26	reaction_26	reaction_26	$S12 \xrightarrow{S12} \emptyset$	
27	reaction_27	reaction_27	$S11 \xrightarrow{S11} \emptyset$	
28	reaction_28	reaction_28	$S12 \xrightarrow{S12} \emptyset$	
29	reaction_29	reaction_29	$S15 \xrightarrow{S15} \emptyset$	
30	reaction_30	reaction_30	$S16 \xrightarrow{S16} S14$	
31	reaction_31	reaction_31	$S15 \xrightarrow{S11, S15} \emptyset$	
32	reaction_32	reaction_32	$S16 \xrightarrow{S11, S16} S14$	
33	reaction_33	reaction_33	$S14 \xrightarrow{S14} \emptyset$	
34	reaction_34	reaction_34	$S16 \xrightarrow{S16} S15$	
35	reaction_35	reaction_35	$S1 + S14 \xrightarrow{S1, S14} S2$	
36	reaction_36	reaction_36	$S14 + S15 \xrightarrow{S14, S15} S16$	
37	reaction_37	reaction_37	$S16 \xrightarrow{S16} S14 + S15$	
38	reaction_38	reaction_38	$S15 \xrightarrow{S18, S15} S21$	

Nº	Id	Name	Reaction Equation	SBO
39	reaction_39	reaction_39	$S_{16} \xrightarrow{S_{18}, S_{16}} S_{14} + S_{21}$	
40	reaction_40	reaction_40	$S_{15} \xrightarrow{S_{18}, S_{15}} S_1$	
41	reaction_41	reaction_41	$S_{16} \xrightarrow{S_{18}, S_{16}} S_2$	
42	reaction_42	reaction_42	$S_{17} \xrightarrow{S_{17}} S_{22}$	
43	reaction_43	reaction_43	$S_{19} \xrightarrow{S_{19}} S_{12}$	
44	reaction_44	reaction_44	$S_{18} \xrightarrow{S_{18}} \emptyset$	
45	reaction_45	reaction_45	$S_{20} \xrightarrow{S_{20}} S_{12}$	
46	reaction_46	reaction_46	$S_{18} \xrightarrow{S_{18}} \emptyset$	
47	reaction_47	reaction_47	$S_{18} \xrightarrow{S_{18}} S_{13}$	
48	reaction_48	reaction_48	$S_{19} \xrightarrow{S_{19}} S_{10} + S_{12}$	
49	reaction_49	reaction_49	$S_{11} + S_{18} \xrightarrow{S_{11}, S_{18}} S_{23}$	
50	reaction_50	reaction_50	$S_{20} \xrightarrow{S_{20}} S_4 + S_{12}$	
51	reaction_51	reaction_51	$S_{19} \xrightarrow{S_{19}} S_{24}$	
52	reaction_52	reaction_52	$S_{20} \xrightarrow{S_{20}} S_3$	
53	reaction_53	reaction_53	$S_{19} \xrightleftharpoons{S_{19}} \emptyset$	
54	reaction_54	reaction_54	$S_{20} \xrightleftharpoons{S_{20}} \emptyset$	
55	reaction_55	reaction_55	$S_{21} \xrightarrow{S_{21}} \emptyset$	
56	reaction_56	reaction_56	$S_{21} \xrightarrow{S_{11}, S_{21}} \emptyset$	
57	reaction_57	reaction_57	$S_{21} \xrightarrow{S_{18}, S_{21}} S_{15}$	
58	reaction_58	reaction_58	$S_{22} \xrightarrow{S_{22}} S_{25}$	
59	reaction_59	reaction_59	$S_{24} \xrightarrow{S_{24}} \emptyset$	

Nº	Id	Name	Reaction Equation	SBO
60	reaction_60	reaction_60	$S_{23} \xrightarrow{S_{23}} S_{11}$	
61	reaction_61	reaction_61	$S_{24} \xrightarrow{S_{24}} S_{11} + S_{26}$	
62	reaction_62	reaction_62	$S_{23} \xrightarrow{S_{23}} S_{11} + S_{18}$	
63	reaction_63	reaction_63	$S_{23} \xrightarrow{S_{18}, S_{14}, S_{23}} S_{18}$	
64	reaction_64	reaction_64	$S_{24} \xrightarrow{S_{18}, S_{14}, S_{24}} S_{26}$	
65	reaction_65	reaction_65	$S_{25} \xrightarrow{S_{25}} S_{27}$	
66	reaction_66	reaction_66	$S_{26} \xrightarrow{S_{26}} \emptyset$	
67	reaction_67	reaction_67	$S_{11} + S_{26} \xrightarrow{S_{11}, S_{26}} S_{24}$	
68	reaction_68	reaction_68	$S_{27} \xrightarrow{S_{27}} S_{28}$	
69	reaction_69	reaction_69	$S_{28} \xrightarrow{S_{28}} \emptyset$	



### 8.1 Reaction `reaction_1`

This is an irreversible reaction of no reactant forming one product.

**Name** `reaction_1`

#### Reaction equation



#### Product

Table 6: Properties of each product.

Id	Name	SBO
S10	@cyto::C4D1(b)	

#### Kinetic Law

**Derived unit** contains undeclared units

$$v_1 = \text{vol}(\text{cell}) \cdot \text{Function\_for\_reaction\_1\_0}(\text{vol}(\text{cell}), \text{ks\_c4}, \text{tf}) \quad (101)$$

$$\text{Function\_for\_reaction\_1\_0}(\text{vol}(\text{cell}), \text{ks\_c4}, \text{tf}) = \frac{\text{ks\_c4} \cdot \text{tf}}{\text{vol}(\text{cell})} \quad (102)$$

$$\text{Function\_for\_reaction\_1\_0}(\text{vol}(\text{cell}), \text{ks\_c4}, \text{tf}) = \frac{\text{ks\_c4} \cdot \text{tf}}{\text{vol}(\text{cell})} \quad (103)$$

### 8.2 Reaction `reaction_2`

This is an irreversible reaction of no reactant forming one product influenced by two modifiers.

**Name** `reaction_2`

#### Reaction equation



#### Modifiers

Table 7: Properties of each modifier.

Id	Name	SBO
S14	@nuc::e2f(b)	
S16	@nuc::e2f(b!1).rb(S788P̃,S800Ũ,b!1)	

## Product

Table 8: Properties of each product.

Id	Name	SBO
S4	@cyto::C2E(T160Ũ,b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_2 = \text{vol}(\text{cell}) \cdot \text{Function\_for\_reaction\_2.0}([S14], [S16], \text{vol}(\text{cell}), \text{ks\_c2e2f}, \text{ks\_c2myc}, \text{tf}) \quad (105)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_2.0}([S14], [S16], \text{vol}(\text{cell}), \text{ks\_c2e2f}, \text{ks\_c2myc}, \text{tf}) \\ &= \frac{\text{ks\_c2myc} \cdot \text{tf} + \text{ks\_c2e2f} \cdot ([S14] + [S16])}{\text{vol}(\text{cell})} \end{aligned} \quad (106)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_2.0}([S14], [S16], \text{vol}(\text{cell}), \text{ks\_c2e2f}, \text{ks\_c2myc}, \text{tf}) \\ &= \frac{\text{ks\_c2myc} \cdot \text{tf} + \text{ks\_c2e2f} \cdot ([S14] + [S16])}{\text{vol}(\text{cell})} \end{aligned} \quad (107)$$

## 8.3 Reaction `reaction_3`

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

**Name** `reaction_3`

### Reaction equation



## Reactant

Table 9: Properties of each reactant.

Id	Name	SBO
S3	@nuc::C2E(T160Û,b!1).p21(b!1)	

## Modifier

Table 10: Properties of each modifier.

Id	Name	SBO
S3	@nuc::C2E(T160Û,b!1).p21(b!1)	

## Product

Table 11: Properties of each product.

Id	Name	SBO
S11	@nuc::p21(b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_3 = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_3\_0}([S3], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \quad (109)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_3\_0}([S3], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \\ &= \frac{(\text{kdeg\_c2} + \text{kdeg\_c2gsk3b} \cdot \text{gsk3b}) \cdot [S3]}{\text{vol}(\text{cell})} \end{aligned} \quad (110)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_3\_0}([S3], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \\ &= \frac{(\text{kdeg\_c2} + \text{kdeg\_c2gsk3b} \cdot \text{gsk3b}) \cdot [S3]}{\text{vol}(\text{cell})} \end{aligned} \quad (111)$$

## 8.4 Reaction `reaction_4`

This is an irreversible reaction of one reactant forming no product influenced by one modifier.

**Name** `reaction_4`

### Reaction equation



### Reactant

Table 12: Properties of each reactant.

Id	Name	SBO
S4	@cyto::C2E(T160Û,b)	

### Modifier

Table 13: Properties of each modifier.

Id	Name	SBO
S4	@cyto::C2E(T160Û,b)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_4 = \text{vol}(\text{cell}) \cdot \text{Function\_for\_reaction\_4\_0}([S4], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \quad (113)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_4\_0}([S4], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \\ &= \frac{(\text{kdeg\_c2} + \text{kdeg\_c2gsk3b} \cdot \text{gsk3b}) \cdot [S4]}{\text{vol}(\text{cell})} \end{aligned} \quad (114)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_4\_0}([S4], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \\ &= \frac{(\text{kdeg\_c2} + \text{kdeg\_c2gsk3b} \cdot \text{gsk3b}) \cdot [S4]}{\text{vol}(\text{cell})} \end{aligned} \quad (115)$$

## 8.5 Reaction `reaction_5`

This is an irreversible reaction of no reactant forming one product influenced by one modifier.

**Name** `reaction_5`

### Reaction equation



## Modifier

Table 14: Properties of each modifier.

Id	Name	SBO
S14	@nuc::e2f(b)	

## Product

Table 15: Properties of each product.

Id	Name	SBO
S12	@cyto::p21(b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_5 = \text{vol}(\text{cell}) \cdot \text{Function\_for\_reaction\_5\_0}([S14], \text{vol}(\text{cell}), \text{ks\_p21e2f}, \text{ks\_p21p53}, \text{tfp21}) \quad (117)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_5\_0}([S14], \text{vol}(\text{cell}), \text{ks\_p21e2f}, \text{ks\_p21p53}, \text{tfp21}) \\ &= \frac{(\text{ks\_p21p53} + \text{ks\_p21e2f} \cdot [S14]) \cdot \text{tfp21}}{\text{vol}(\text{cell})} \end{aligned} \quad (118)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_5\_0}([S14], \text{vol}(\text{cell}), \text{ks\_p21e2f}, \text{ks\_p21p53}, \text{tfp21}) \\ &= \frac{(\text{ks\_p21p53} + \text{ks\_p21e2f} \cdot [S14]) \cdot \text{tfp21}}{\text{vol}(\text{cell})} \end{aligned} \quad (119)$$

## 8.6 Reaction `reaction_6`

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

**Name** `reaction_6`

### Reaction equation



## Reactant

Table 16: Properties of each reactant.

Id	Name	SBO
S3	@nuc::C2E(T160 $\tilde{U}$ ,b!1).p21(b!1)	

## Modifier

Table 17: Properties of each modifier.

Id	Name	SBO
S3	@nuc::C2E(T160 $\tilde{U}$ ,b!1).p21(b!1)	

## Products

Table 18: Properties of each product.

Id	Name	SBO
S11	@nuc::p21(b)	
S13	@nuc::C2E(T160 $\tilde{U}$ ,b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_6 = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_6\_0}([S3], \text{vol}(\text{cell}), \text{kd\_p21c2}) \quad (121)$$

$$\text{Function\_for\_reaction\_6\_0}([S3], \text{vol}(\text{cell}), \text{kd\_p21c2}) = \frac{\text{kd\_p21c2} \cdot [S3]}{\text{vol}(\text{cell})} \quad (122)$$

$$\text{Function\_for\_reaction\_6\_0}([S3], \text{vol}(\text{cell}), \text{kd\_p21c2}) = \frac{\text{kd\_p21c2} \cdot [S3]}{\text{vol}(\text{cell})} \quad (123)$$

## 8.7 Reaction `reaction_7`

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

**Name** `reaction_7`

### Reaction equation



## Reactant

Table 19: Properties of each reactant.

Id	Name	SBO
S3	@nuc::C2E(T160 $\tilde{U}$ ,b!1).p21(b!1)	

## Modifiers

Table 20: Properties of each modifier.

Id	Name	SBO
S18	@nuc::C2E(T160 $\tilde{P}$ ,b)	
S14	@nuc::e2f(b)	
S3	@nuc::C2E(T160 $\tilde{U}$ ,b!1).p21(b!1)	

## Product

Table 21: Properties of each product.

Id	Name	SBO
S13	@nuc::C2E(T160 $\tilde{U}$ ,b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_7 = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_7\_0}([S14], [S18], [S3], \text{vol}(\text{cell}), \text{erk}, \text{kdeg\_p21c2skp2}, \text{kdeg\_p21erkskp2}, \text{kdeg\_p21skp2}) \quad (125)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_7\_0}([S14], [S18], [S3], \text{vol}(\text{cell}), \text{erk}, \\ & \text{kdeg\_p21c2skp2}, \text{kdeg\_p21erkskp2}, \text{kdeg\_p21skp2}) \\ &= \frac{(\text{kdeg\_p21erkskp2} \cdot \text{erk} + \text{kdeg\_p21c2skp2} \cdot [S18] + \text{kdeg\_p21skp2}) \cdot [S14] \cdot [S3]}{\text{vol}(\text{cell})} \end{aligned} \quad (126)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_7\_0}([S14], [S18], [S3], \text{vol}(\text{cell}), \text{erk}, \\ & \text{kdeg\_p21c2skp2}, \text{kdeg\_p21erkskp2}, \text{kdeg\_p21skp2}) \\ &= \frac{(\text{kdeg\_p21erkskp2} \cdot \text{erk} + \text{kdeg\_p21c2skp2} \cdot [S18] + \text{kdeg\_p21skp2}) \cdot [S14] \cdot [S3]}{\text{vol}(\text{cell})} \end{aligned} \quad (127)$$

## 8.8 Reaction `reaction_8`

This is an irreversible reaction of no reactant forming one product influenced by one modifier.

**Name** `reaction_8`

### Reaction equation



### Modifier

Table 22: Properties of each modifier.

Id	Name	SBO
S14	@nuc::e2f(b)	

### Product

Table 23: Properties of each product.

Id	Name	SBO
S1	@nuc::rb(S788Û,S800Û,b)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_8 = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_8\_0}([S14], \text{vol}(\text{cell}), ks\_rb, ks\_rbe2f) \quad (129)$$

$$\text{Function\_for\_reaction\_8\_0}([S14], \text{vol}(\text{cell}), ks\_rb, ks\_rbe2f) = \frac{ks\_rb + ks\_rbe2f \cdot [S14]}{\text{vol}(\text{cell})} \quad (130)$$

$$\text{Function\_for\_reaction\_8\_0}([S14], \text{vol}(\text{cell}), ks\_rb, ks\_rbe2f) = \frac{ks\_rb + ks\_rbe2f \cdot [S14]}{\text{vol}(\text{cell})} \quad (131)$$

## 8.9 Reaction `reaction_9`

This is an irreversible reaction of one reactant forming no product influenced by one modifier.

**Name** `reaction_9`



### Reaction equation



### Reactant

Table 24: Properties of each reactant.

Id	Name	SBO
S1	@nuc::rb(S788Û,S800Û,b)	

### Modifier

Table 25: Properties of each modifier.

Id	Name	SBO
S1	@nuc::rb(S788Û,S800Û,b)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_9 = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_9\_0}([S1], \text{vol}(\text{cell}), \text{kdeg\_rbfree}) \quad (133)$$

$$\text{Function\_for\_reaction\_9\_0}([S1], \text{vol}(\text{cell}), \text{kdeg\_rbfree}) = \frac{\text{kdeg\_rbfree} \cdot [S1]}{\text{vol}(\text{cell})} \quad (134)$$

$$\text{Function\_for\_reaction\_9\_0}([S1], \text{vol}(\text{cell}), \text{kdeg\_rbfree}) = \frac{\text{kdeg\_rbfree} \cdot [S1]}{\text{vol}(\text{cell})} \quad (135)$$

## 8.10 Reaction `reaction_10`

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

**Name** `reaction_10`

### Reaction equation



### Reactant

Table 26: Properties of each reactant.

Id	Name	SBO
S2	@nuc::e2f(b!1).rb(S788Û,S800Û,b!1)	

## Modifier

Table 27: Properties of each modifier.

Id	Name	SBO
S2	@nuc::e2f(b!1).rb(S788Û,S800Û,b!1)	

## Product

Table 28: Properties of each product.

Id	Name	SBO
S14	@nuc::e2f(b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{10} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_10\_0}([S2], \text{vol}(\text{cell}), \text{kdeg\_rbbound}) \quad (137)$$

$$\text{Function\_for\_reaction\_10\_0}([S2], \text{vol}(\text{cell}), \text{kdeg\_rbbound}) = \frac{\text{kdeg\_rbbound} \cdot [S2]}{\text{vol}(\text{cell})} \quad (138)$$

$$\text{Function\_for\_reaction\_10\_0}([S2], \text{vol}(\text{cell}), \text{kdeg\_rbbound}) = \frac{\text{kdeg\_rbbound} \cdot [S2]}{\text{vol}(\text{cell})} \quad (139)$$

### 8.11 Reaction `reaction_11`

This is an irreversible reaction of one reactant forming no product influenced by two modifiers.

**Name** `reaction_11`

### Reaction equation



## Reactant

Table 29: Properties of each reactant.

Id	Name	SBO
S1	@nuc::rb(S788 $\tilde{U}$ ,S800 $\tilde{U}$ ,b)	

## Modifiers

Table 30: Properties of each modifier.

Id	Name	SBO
S11	@nuc::p21(b)	
S1	@nuc::rb(S788 $\tilde{U}$ ,S800 $\tilde{U}$ ,b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{11} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_11\_0}([S1], [S11], \text{vol}(\text{cell}), \text{kdeg\_rbp21}) \quad (141)$$

$$\text{Function\_for\_reaction\_11\_0}([S1], [S11], \text{vol}(\text{cell}), \text{kdeg\_rbp21}) = \frac{\text{kdeg\_rbp21} \cdot [S11] \cdot [S1]}{\text{vol}(\text{cell})} \quad (142)$$

$$\text{Function\_for\_reaction\_11\_0}([S1], [S11], \text{vol}(\text{cell}), \text{kdeg\_rbp21}) = \frac{\text{kdeg\_rbp21} \cdot [S11] \cdot [S1]}{\text{vol}(\text{cell})} \quad (143)$$

## 8.12 Reaction `reaction_12`

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

**Name** `reaction_12`

### Reaction equation



### Reactant

Table 31: Properties of each reactant.

Id	Name	SBO
S2	@nuc::e2f(b!1).rb(S788Û,S800Û,b!1)	

## Modifiers

Table 32: Properties of each modifier.

Id	Name	SBO
S11	@nuc::p21(b)	
S2	@nuc::e2f(b!1).rb(S788Û,S800Û,b!1)	

## Product

Table 33: Properties of each product.

Id	Name	SBO
S14	@nuc::e2f(b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{12} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_12\_0}([S11], [S2], \text{vol}(\text{cell}), \text{kdeg\_rbp21}) \quad (145)$$

$$\text{Function\_for\_reaction\_12\_0}([S11], [S2], \text{vol}(\text{cell}), \text{kdeg\_rbp21}) = \frac{\text{kdeg\_rbp21} \cdot [S11] \cdot [S2]}{\text{vol}(\text{cell})} \quad (146)$$

$$\text{Function\_for\_reaction\_12\_0}([S11], [S2], \text{vol}(\text{cell}), \text{kdeg\_rbp21}) = \frac{\text{kdeg\_rbp21} \cdot [S11] \cdot [S2]}{\text{vol}(\text{cell})} \quad (147)$$

### 8.13 Reaction `reaction_13`

This is an irreversible reaction of no reactant forming one product influenced by one modifier.

**Name** `reaction_13`

#### Reaction equation



## Modifier

Table 34: Properties of each modifier.

Id	Name	SBO
S14	@nuc::e2f(b)	

## Product

Table 35: Properties of each product.

Id	Name	SBO
S14	@nuc::e2f(b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{13} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_13\_0}([S14], \text{vol}(\text{cell}), \text{ks\_e2fe2f}, \text{ks\_e2fmyc}, \text{tf}) \quad (149)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_13\_0}([S14], \text{vol}(\text{cell}), \text{ks\_e2fe2f}, \text{ks\_e2fmyc}, \text{tf}) \\ &= \frac{(\text{ks\_e2fe2f} \cdot [S14] + \text{ks\_e2fmyc}) \cdot \text{tf}}{\text{vol}(\text{cell})} \end{aligned} \quad (150)$$

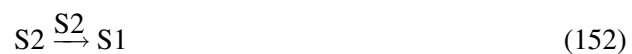
$$\begin{aligned} & \text{Function\_for\_reaction\_13\_0}([S14], \text{vol}(\text{cell}), \text{ks\_e2fe2f}, \text{ks\_e2fmyc}, \text{tf}) \\ &= \frac{(\text{ks\_e2fe2f} \cdot [S14] + \text{ks\_e2fmyc}) \cdot \text{tf}}{\text{vol}(\text{cell})} \end{aligned} \quad (151)$$

### 8.14 Reaction `reaction_14`

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

**Name** `reaction_14`

#### Reaction equation



## Reactant

Table 36: Properties of each reactant.

Id	Name	SBO
S2	@nuc::e2f(b!1).rb(S788Û,S800Û,b!1)	

## Modifier

Table 37: Properties of each modifier.

Id	Name	SBO
S2	@nuc::e2f(b!1).rb(S788Û,S800Û,b!1)	

## Product

Table 38: Properties of each product.

Id	Name	SBO
S1	@nuc::rb(S788Û,S800Û,b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{14} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_14\_0}([S2], \text{vol}(\text{cell}), \text{kdeg\_e2fbound}) \quad (153)$$

$$\text{Function\_for\_reaction\_14\_0}([S2], \text{vol}(\text{cell}), \text{kdeg\_e2fbound}) = \frac{\text{kdeg\_e2fbound} \cdot [S2]}{\text{vol}(\text{cell})} \quad (154)$$

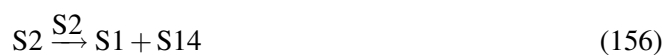
$$\text{Function\_for\_reaction\_14\_0}([S2], \text{vol}(\text{cell}), \text{kdeg\_e2fbound}) = \frac{\text{kdeg\_e2fbound} \cdot [S2]}{\text{vol}(\text{cell})} \quad (155)$$

### 8.15 Reaction `reaction_15`

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

**Name** `reaction_15`

#### Reaction equation



## Reactant

Table 39: Properties of each reactant.

Id	Name	SBO
S2	@nuc::e2f(b!1).rb(S788Û,S800Û,b!1)	

## Modifier

Table 40: Properties of each modifier.

Id	Name	SBO
S2	@nuc::e2f(b!1).rb(S788Û,S800Û,b!1)	

## Products

Table 41: Properties of each product.

Id	Name	SBO
S1	@nuc::rb(S788Û,S800Û,b)	
S14	@nuc::e2f(b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{15} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_15\_0}([S2], \text{vol}(\text{cell}), \text{kd\_rbe2f}) \quad (157)$$

$$\text{Function\_for\_reaction\_15\_0}([S2], \text{vol}(\text{cell}), \text{kd\_rbe2f}) = \frac{\text{kd\_rbe2f} \cdot [S2]}{\text{vol}(\text{cell})} \quad (158)$$

$$\text{Function\_for\_reaction\_15\_0}([S2], \text{vol}(\text{cell}), \text{kd\_rbe2f}) = \frac{\text{kd\_rbe2f} \cdot [S2]}{\text{vol}(\text{cell})} \quad (159)$$

## 8.16 Reaction `reaction_16`

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

**Name** `reaction_16`

### Reaction equation



## Reactant

Table 42: Properties of each reactant.

Id	Name	SBO
S1	@nuc::rb(S788 $\tilde{U}$ ,S800 $\tilde{U}$ ,b)	

## Modifiers

Table 43: Properties of each modifier.

Id	Name	SBO
S24	@nuc::C4D1(b!1),p21(b!1)	
S1	@nuc::rb(S788 $\tilde{U}$ ,S800 $\tilde{U}$ ,b)	

## Product

Table 44: Properties of each product.

Id	Name	SBO
S15	@nuc::rb(S788 $\tilde{P}$ ,S800 $\tilde{U}$ ,b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{16} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_16\_0}(\text{Km\_prb}, [\text{S1}], [\text{S24}], \text{vol}(\text{cell}), \text{kcatp\_rbc4}, \text{nrb}) \quad (161)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_16\_0}(\text{Km\_prb}, [\text{S1}], [\text{S24}], \text{vol}(\text{cell}), \text{kcatp\_rbc4}, \text{nrb}) \\ &= \frac{\frac{\text{kcatp\_rbc4} \cdot [\text{S24}] \cdot [\text{S1}]^{\text{nrb}}}{\text{Km\_prb}^{\text{nrb}} + [\text{S1}]^{\text{nrb}}}}{\text{vol}(\text{cell})} \end{aligned} \quad (162)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_16\_0}(\text{Km\_prb}, [\text{S1}], [\text{S24}], \text{vol}(\text{cell}), \text{kcatp\_rbc4}, \text{nrb}) \\ &= \frac{\frac{\text{kcatp\_rbc4} \cdot [\text{S24}] \cdot [\text{S1}]^{\text{nrb}}}{\text{Km\_prb}^{\text{nrb}} + [\text{S1}]^{\text{nrb}}}}{\text{vol}(\text{cell})} \end{aligned} \quad (163)$$

### 8.17 Reaction `reaction_17`

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

**Name** `reaction_17`



## Reaction equation



## Reactant

Table 45: Properties of each reactant.

Id	Name	SBO
S2	@nuc::e2f(b!1).rb(S788Û,S800Û,b!1)	

## Modifiers

Table 46: Properties of each modifier.

Id	Name	SBO
S24	@nuc::C4D1(b!1),p21(b!1)	
S2	@nuc::e2f(b!1).rb(S788Û,S800Û,b!1)	

## Product

Table 47: Properties of each product.

Id	Name	SBO
S16	@nuc::e2f(b!1).rb(S788P̃,S800Û,b!1)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{17} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_17\_0}(\text{Km\_prb}, [S2], [S24], \text{vol}(\text{cell}), \text{kcatp\_rbc4}, \text{nr}) \quad (165)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_17\_0}(\text{Km\_prb}, [S2], [S24], \text{vol}(\text{cell}), \text{kcatp\_rbc4}, \text{nr}) \\ &= \frac{\frac{\text{kcatp\_rbc4} \cdot [S24] \cdot [S2]^{\text{nr}}}{\text{Km\_prb}^{\text{nr}} + [S2]^{\text{nr}}}}{\text{vol}(\text{cell})} \end{aligned} \quad (166)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_17\_0}(\text{Km\_prb}, [S2], [S24], \text{vol}(\text{cell}), \text{kcatp\_rbc4}, \text{nr}) \\ &= \frac{\frac{\text{kcatp\_rbc4} \cdot [S24] \cdot [S2]^{\text{nr}}}{\text{Km\_prb}^{\text{nr}} + [S2]^{\text{nr}}}}{\text{vol}(\text{cell})} \end{aligned} \quad (167)$$

### 8.18 Reaction `reaction_18`

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

**Name** `reaction_18`

#### Reaction equation



#### Reactant

Table 48: Properties of each reactant.

Id	Name	SBO
S5	@nuc::dnapre()	

#### Modifiers

Table 49: Properties of each modifier.

Id	Name	SBO
S18	@nuc::C2E(T160P̃,b)	
S14	@nuc::e2f(b)	
S5	@nuc::dnapre()	

#### Product

Table 50: Properties of each product.

Id	Name	SBO
S17	@nuc::dnapre1()	

#### Kinetic Law

**Derived unit** contains undeclared units

$$v_{18} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_18\_0}([S14], [S18], [S5], \text{vol}(\text{cell}), k_{\text{dna}}) \quad (169)$$

$$\text{Function\_for\_reaction\_18\_0}([S14], [S18], [S5], \text{vol}(\text{cell}), k_{\text{dna}}) = \frac{k_{\text{dna}} \cdot [S18] \cdot [S14] \cdot [S5]}{\text{vol}(\text{cell})} \quad (170)$$

$$\text{Function\_for\_reaction\_18\_0}([S14], [S18], [S5], \text{vol}(\text{cell}), k_{\text{dna}}) = \frac{k_{\text{dna}} \cdot [S18] \cdot [S14] \cdot [S5]}{\text{vol}(\text{cell})} \quad (171)$$

### 8.19 Reaction `reaction_19`

This is an irreversible reaction of one reactant forming no product influenced by one modifier.

**Name** `reaction_19`

#### Reaction equation



#### Reactant

Table 51: Properties of each reactant.

Id	Name	SBO
S10	@cyto::C4D1(b)	

#### Modifier

Table 52: Properties of each modifier.

Id	Name	SBO
S10	@cyto::C4D1(b)	

#### Kinetic Law

**Derived unit** contains undeclared units

$$v_{19} = \text{vol}(\text{cell}) \cdot \text{Function\_for\_reaction\_19\_0}([S10], \text{vol}(\text{cell}), \text{gsk3b}, k_{\text{deg\_c4}}, k_{\text{deg\_c4gsk3b}}) \quad (173)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_19\_0}([S10], \text{vol}(\text{cell}), \text{gsk3b}, k_{\text{deg\_c4}}, k_{\text{deg\_c4gsk3b}}) \\ &= \frac{(k_{\text{deg\_c4}} + k_{\text{deg\_c4gsk3b}} \cdot \text{gsk3b}) \cdot [S10]}{\text{vol}(\text{cell})} \end{aligned} \quad (174)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_19\_0}([S10], \text{vol}(\text{cell}), \text{gsk3b}, k_{\text{deg\_c4}}, k_{\text{deg\_c4gsk3b}}) \\ &= \frac{(k_{\text{deg\_c4}} + k_{\text{deg\_c4gsk3b}} \cdot \text{gsk3b}) \cdot [S10]}{\text{vol}(\text{cell})} \end{aligned} \quad (175)$$

## 8.20 Reaction `reaction_20`

This is an irreversible reaction of one reactant forming no product influenced by one modifier.

**Name** `reaction_20`

### Reaction equation



### Reactant

Table 53: Properties of each reactant.

Id	Name	SBO
S13	@nuc::C2E(T160Û,b)	

### Modifier

Table 54: Properties of each modifier.

Id	Name	SBO
S13	@nuc::C2E(T160Û,b)	

### Kinetic Law

**Derived unit** contains undeclared units

$$\begin{aligned} v_{20} &= \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_20\_0}([S13], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \\ &\quad (177) \end{aligned}$$

$$\begin{aligned} &\text{Function\_for\_reaction\_20\_0}([S13], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \\ &= \frac{(\text{kdeg\_c2} + \text{kdeg\_c2gsk3b} \cdot \text{gsk3b}) \cdot [S13]}{\text{vol}(\text{cell})} \quad (178) \end{aligned}$$

$$\begin{aligned} &\text{Function\_for\_reaction\_20\_0}([S13], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \\ &= \frac{(\text{kdeg\_c2} + \text{kdeg\_c2gsk3b} \cdot \text{gsk3b}) \cdot [S13]}{\text{vol}(\text{cell})} \quad (179) \end{aligned}$$

## 8.21 Reaction `reaction_21`

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

**Name** `reaction_21`

### Reaction equation



### Reactant

Table 55: Properties of each reactant.

Id	Name	SBO
S13	@nuc::C2E(T160Û,b)	

### Modifier

Table 56: Properties of each modifier.

Id	Name	SBO
S13	@nuc::C2E(T160Û,b)	

### Product

Table 57: Properties of each product.

Id	Name	SBO
S18	@nuc::C2E(T160P̃,b)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{21} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_21\_0}([S13], \text{vol}(\text{cell}), \text{kp\_c2cak}) \quad (181)$$

$$\text{Function\_for\_reaction\_21\_0}([S13], \text{vol}(\text{cell}), \text{kp\_c2cak}) = \frac{\text{kp\_c2cak} \cdot [S13]}{\text{vol}(\text{cell})} \quad (182)$$

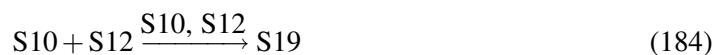
$$\text{Function\_for\_reaction\_21\_0}([S13], \text{vol}(\text{cell}), \text{kp\_c2cak}) = \frac{\text{kp\_c2cak} \cdot [S13]}{\text{vol}(\text{cell})} \quad (183)$$

## 8.22 Reaction `reaction_22`

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

**Name** `reaction_22`

### Reaction equation



### Reactants

Table 58: Properties of each reactant.

Id	Name	SBO
S10	@cyto::C4D1(b)	
S12	@cyto::p21(b)	

### Modifiers

Table 59: Properties of each modifier.

Id	Name	SBO
S10	@cyto::C4D1(b)	
S12	@cyto::p21(b)	

### Product

Table 60: Properties of each product.

Id	Name	SBO
S19	@cyto::C4D1(b!1).p21(b!1)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{22} = \text{vol}(\text{cell}) \cdot \text{Function\_for\_reaction\_22\_0}([S10], [S12], \text{vol}(\text{cell}), \text{kb\_p21c4}) \quad (185)$$

$$\text{Function\_for\_reaction\_22\_0}([S10], [S12], \text{vol}(\text{cell}), \text{kb\_p21c4}) = \frac{\text{kb\_p21c4} \cdot [S10] \cdot [S12]}{\text{vol}(\text{cell})} \quad (186)$$

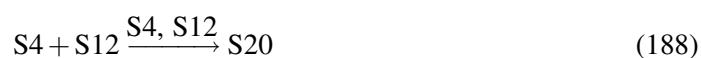
$$\text{Function\_for\_reaction\_22\_0}([S10], [S12], \text{vol}(\text{cell}), \text{kb\_p21c4}) = \frac{\text{kb\_p21c4} \cdot [S10] \cdot [S12]}{\text{vol}(\text{cell})} \quad (187)$$

### 8.23 Reaction `reaction_23`

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

**Name** `reaction_23`

#### Reaction equation



#### Reactants

Table 61: Properties of each reactant.

Id	Name	SBO
S4	@cyto::C2E(T160Û,b)	
S12	@cyto::p21(b)	

#### Modifiers

Table 62: Properties of each modifier.

Id	Name	SBO
S4	@cyto::C2E(T160Û,b)	
S12	@cyto::p21(b)	

#### Product

Table 63: Properties of each product.

Id	Name	SBO
S20	@cyto::C2E(T160Û,b!1).p21(b!1)	

#### Kinetic Law

**Derived unit** contains undeclared units

$$v_{23} = \text{vol}(\text{cell}) \cdot \text{Function\_for\_reaction\_23\_0}([S12], [S4], \text{vol}(\text{cell}), \text{kb\_p21c2}) \quad (189)$$

$$\text{Function\_for\_reaction\_23\_0}([S12],[S4],\text{vol}(\text{cell}),\text{kb\_p21c2}) = \frac{\text{kb\_p21c2} \cdot [S4] \cdot [S12]}{\text{vol}(\text{cell})} \quad (190)$$

$$\text{Function\_for\_reaction\_23\_0}([S12],[S4],\text{vol}(\text{cell}),\text{kb\_p21c2}) = \frac{\text{kb\_p21c2} \cdot [S4] \cdot [S12]}{\text{vol}(\text{cell})} \quad (191)$$

## 8.24 Reaction `reaction_24`

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

**Name** `reaction_24`

### Reaction equation



### Reactants

Table 64: Properties of each reactant.

Id	Name	SBO
S11	@nuc::p21(b)	
S13	@nuc::C2E(T160Û,b)	

### Modifiers

Table 65: Properties of each modifier.

Id	Name	SBO
S11	@nuc::p21(b)	
S13	@nuc::C2E(T160Û,b)	

### Product

Table 66: Properties of each product.

Id	Name	SBO
S3	@nuc::C2E(T160Û,b!1).p21(b!1)	



## Kinetic Law

**Derived unit** contains undeclared units

$$v_{24} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_24\_0}([S11], [S13], \text{vol}(\text{cell}), \text{kb\_p21c2}) \quad (193)$$

$$\text{Function\_for\_reaction\_24\_0}([S11], [S13], \text{vol}(\text{cell}), \text{kb\_p21c2}) = \frac{\text{kb\_p21c2} \cdot [S11] \cdot [S13]}{\text{vol}(\text{cell})} \quad (194)$$

$$\text{Function\_for\_reaction\_24\_0}([S11], [S13], \text{vol}(\text{cell}), \text{kb\_p21c2}) = \frac{\text{kb\_p21c2} \cdot [S11] \cdot [S13]}{\text{vol}(\text{cell})} \quad (195)$$

## 8.25 Reaction `reaction_25`

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

**Name** `reaction_25`

### Reaction equation



### Reactant

Table 67: Properties of each reactant.

Id	Name	SBO
S12	@cyto::p21(b)	

### Modifier

Table 68: Properties of each modifier.

Id	Name	SBO
S12	@cyto::p21(b)	

### Product

Table 69: Properties of each product.

Id	Name	SBO
S11	@nuc::p21(b)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{25} = \text{Function\_for\_reaction\_25\_0}([S12], \text{Vratio}, \text{vol}(\text{cell}), \text{kimport}) \quad (197)$$

$$\text{Function\_for\_reaction\_25\_0}([S12], \text{Vratio}, \text{vol}(\text{cell}), \text{kimport}) = \frac{\frac{\text{kimport}}{\text{Vratio}} \cdot [S12]}{\text{vol}(\text{cell})} \quad (198)$$

## 8.26 Reaction reaction\_26

This is a reversible reaction of one reactant forming no product influenced by one modifier.

**Name** reaction\_26

### Reaction equation



### Reactant

Table 70: Properties of each reactant.

Id	Name	SBO
S12	@cyto::p21(b)	

### Modifier

Table 71: Properties of each modifier.

Id	Name	SBO
S12	@cyto::p21(b)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{26} = \text{vol}(\text{cell}) \cdot \text{Function\_for\_reaction\_26\_0}([S12], V_{\text{ratio}}, \text{vol}(\text{cell}), \text{kimport}) \quad (200)$$

$$\text{Function\_for\_reaction\_26\_0}([S12], V_{\text{ratio}}, \text{vol}(\text{cell}), \text{kimport}) = \frac{\text{kimport} \cdot \left(1 - \frac{1}{V_{\text{ratio}}}\right) \cdot [S12]}{\text{vol}(\text{cell})} \quad (201)$$

$$\text{Function\_for\_reaction\_26\_0}([S12], V_{\text{ratio}}, \text{vol}(\text{cell}), \text{kimport}) = \frac{\text{kimport} \cdot \left(1 - \frac{1}{V_{\text{ratio}}}\right) \cdot [S12]}{\text{vol}(\text{cell})} \quad (202)$$

## 8.27 Reaction `reaction_27`

This is an irreversible reaction of one reactant forming no product influenced by one modifier.

**Name** `reaction_27`

### Reaction equation



### Reactant

Table 72: Properties of each reactant.

Id	Name	SBO
S11	@nuc::p21(b)	

### Modifier

Table 73: Properties of each modifier.

Id	Name	SBO
S11	@nuc::p21(b)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{27} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_27\_0}([S11], \text{vol}(\text{cell}), \text{erk}, \text{gsk3b}, \text{kdeg\_p21erk}, \text{kdeg\_p21gsk3b}) \quad (204)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_27\_0}([S11], \text{vol}(\text{cell}), \text{erk}, \text{gsk3b}, \text{kdeg\_p21erk}, \text{kdeg\_p21gsk3b}) \\ &= \frac{(\text{kdeg\_p21gsk3b} \cdot \text{gsk3b} + \text{kdeg\_p21erk} \cdot \text{erk}) \cdot [S11]}{\text{vol}(\text{cell})} \end{aligned} \quad (205)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_27\_0}([S11], \text{vol}(\text{cell}), \text{erk}, \text{gsk3b}, \text{kdeg\_p21erk}, \text{kdeg\_p21gsk3b}) \\ &= \frac{(\text{kdeg\_p21gsk3b} \cdot \text{gsk3b} + \text{kdeg\_p21erk} \cdot \text{erk}) \cdot [S11]}{\text{vol}(\text{cell})} \end{aligned} \quad (206)$$

## 8.28 Reaction `reaction_28`

This is an irreversible reaction of one reactant forming no product influenced by one modifier.

**Name** `reaction_28`

### Reaction equation



### Reactant

Table 74: Properties of each reactant.

Id	Name	SBO
S12	@cyto::p21(b)	

### Modifier

Table 75: Properties of each modifier.

Id	Name	SBO
S12	@cyto::p21(b)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{28} = \text{vol}(\text{cell}) \cdot \text{Function\_for\_reaction\_28\_0}([S12], \text{vol}(\text{cell}), \text{erk}, \text{gsk3b}, \text{kdeg\_p21erk}, \text{kdeg\_p21gsk3b}) \quad (208)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_28\_0}([S12], \text{vol}(\text{cell}), \text{erk}, \text{gsk3b}, \text{kdeg\_p21erk}, \text{kdeg\_p21gsk3b}) \\ &= \frac{(\text{kdeg\_p21gsk3b} \cdot \text{gsk3b} + \text{kdeg\_p21erk} \cdot \text{erk}) \cdot [S12]}{\text{vol}(\text{cell})} \end{aligned} \quad (209)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_28\_0}([S12], \text{vol}(\text{cell}), \text{erk}, \text{gsk3b}, \text{kdeg\_p21erk}, \text{kdeg\_p21gsk3b}) \\ &= \frac{(\text{kdeg\_p21gsk3b} \cdot \text{gsk3b} + \text{kdeg\_p21erk} \cdot \text{erk}) \cdot [S12]}{\text{vol}(\text{cell})} \end{aligned} \quad (210)$$

## 8.29 Reaction `reaction_29`

This is an irreversible reaction of one reactant forming no product influenced by one modifier.

**Name** `reaction_29`

### Reaction equation



### Reactant

Table 76: Properties of each reactant.

Id	Name	SBO
S15	@nuc::rb(S788P̃,S800Ũ,b)	

### Modifier

Table 77: Properties of each modifier.

Id	Name	SBO
S15	@nuc::rb(S788P̃,S800Ũ,b)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{29} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_29\_0}([S15], \text{vol}(\text{cell}), \text{kdeg\_rbfree}) \quad (212)$$

$$\text{Function\_for\_reaction\_29\_0}([S15], \text{vol}(\text{cell}), \text{kdeg\_rbfree}) = \frac{\text{kdeg\_rbfree} \cdot [S15]}{\text{vol}(\text{cell})} \quad (213)$$

$$\text{Function\_for\_reaction\_29\_0}([S15], \text{vol}(\text{cell}), \text{kdeg\_rbfree}) = \frac{\text{kdeg\_rbfree} \cdot [S15]}{\text{vol}(\text{cell})} \quad (214)$$

### 8.30 Reaction `reaction_30`

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

**Name** `reaction_30`

#### Reaction equation



#### Reactant

Table 78: Properties of each reactant.

Id	Name	SBO
S16	@nuc::e2f(b!1).rb(S788P̃,S800Ũ,b!1)	

#### Modifier

Table 79: Properties of each modifier.

Id	Name	SBO
S16	@nuc::e2f(b!1).rb(S788P̃,S800Ũ,b!1)	

#### Product

Table 80: Properties of each product.

Id	Name	SBO
S14	@nuc::e2f(b)	

#### Kinetic Law

**Derived unit** contains undeclared units

$$v_{30} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_30\_0}([S16], \text{vol}(\text{cell}), \text{kdeg\_rbbound}) \quad (216)$$

$$\text{Function\_for\_reaction\_30\_0}([S16], \text{vol}(\text{cell}), \text{kdeg\_rbbound}) = \frac{\text{kdeg\_rbbound} \cdot [S16]}{\text{vol}(\text{cell})} \quad (217)$$

$$\text{Function\_for\_reaction\_30\_0}([S16], \text{vol}(\text{cell}), \text{kdeg\_rbbound}) = \frac{\text{kdeg\_rbbound} \cdot [S16]}{\text{vol}(\text{cell})} \quad (218)$$

### 8.31 Reaction `reaction_31`

This is an irreversible reaction of one reactant forming no product influenced by two modifiers.

**Name** `reaction_31`

#### Reaction equation



#### Reactant

Table 81: Properties of each reactant.

Id	Name	SBO
S15	@nuc::rb(S788P̃,S800Ũ,b)	

#### Modifiers

Table 82: Properties of each modifier.

Id	Name	SBO
S11	@nuc::p21(b)	
S15	@nuc::rb(S788P̃,S800Ũ,b)	

#### Kinetic Law

**Derived unit** contains undeclared units

$$v_{31} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_31\_0}([S11], [S15], \text{vol}(\text{cell}), \text{kdeg\_rbp21}) \quad (220)$$

$$\text{Function\_for\_reaction\_31\_0}([S11], [S15], \text{vol}(\text{cell}), \text{kdeg\_rbp21}) = \frac{\text{kdeg\_rbp21} \cdot [S11] \cdot [S15]}{\text{vol}(\text{cell})} \quad (221)$$

$$\text{Function\_for\_reaction\_31\_0}([S11], [S15], \text{vol}(\text{cell}), \text{kdeg\_rbp21}) = \frac{\text{kdeg\_rbp21} \cdot [S11] \cdot [S15]}{\text{vol}(\text{cell})} \quad (222)$$

### 8.32 Reaction `reaction_32`

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

**Name** `reaction_32`

#### Reaction equation



#### Reactant

Table 83: Properties of each reactant.

Id	Name	SBO
S16	@nuc::e2f(b!1).rb(S788P̃,S800Ũ,b!1)	

#### Modifiers

Table 84: Properties of each modifier.

Id	Name	SBO
S11	@nuc::p21(b)	
S16	@nuc::e2f(b!1).rb(S788P̃,S800Ũ,b!1)	

#### Product

Table 85: Properties of each product.

Id	Name	SBO
S14	@nuc::e2f(b)	

#### Kinetic Law

**Derived unit** contains undeclared units

$$v_{32} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_32\_0}([S11], [S16], \text{vol}(\text{cell}), \text{kdeg\_rbp21}) \quad (224)$$

$$\text{Function\_for\_reaction\_32\_0}([S11], [S16], \text{vol}(\text{cell}), \text{kdeg\_rbp21}) = \frac{\text{kdeg\_rbp21} \cdot [S11] \cdot [S16]}{\text{vol}(\text{cell})} \quad (225)$$



$$\text{Function\_for\_reaction\_32\_0}([S11],[S16],\text{vol}(\text{cell}),\text{kdeg\_rbp21}) = \frac{\text{kdeg\_rbp21} \cdot [S11] \cdot [S16]}{\text{vol}(\text{cell})} \quad (226)$$

### 8.33 Reaction `reaction_33`

This is an irreversible reaction of one reactant forming no product influenced by one modifier.

**Name** `reaction_33`

#### Reaction equation



#### Reactant

Table 86: Properties of each reactant.

Id	Name	SBO
S14	@nuc::e2f(b)	

#### Modifier

Table 87: Properties of each modifier.

Id	Name	SBO
S14	@nuc::e2f(b)	

#### Kinetic Law

**Derived unit** contains undeclared units

$$v_{33} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_33\_0}([S14],\text{vol}(\text{cell}),\text{kdeg\_e2ffree}) \quad (228)$$

$$\text{Function\_for\_reaction\_33\_0}([S14],\text{vol}(\text{cell}),\text{kdeg\_e2ffree}) = \frac{\text{kdeg\_e2ffree} \cdot [S14]}{\text{vol}(\text{cell})} \quad (229)$$

$$\text{Function\_for\_reaction\_33\_0}([S14],\text{vol}(\text{cell}),\text{kdeg\_e2ffree}) = \frac{\text{kdeg\_e2ffree} \cdot [S14]}{\text{vol}(\text{cell})} \quad (230)$$

### 8.34 Reaction `reaction_34`

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

**Name** `reaction_34`

## Reaction equation



## Reactant

Table 88: Properties of each reactant.

Id	Name	SBO
S16	@nuc::e2f(b!1).rb(S788P̃,S800Ũ,b!1)	

## Modifier

Table 89: Properties of each modifier.

Id	Name	SBO
S16	@nuc::e2f(b!1).rb(S788P̃,S800Ũ,b!1)	

## Product

Table 90: Properties of each product.

Id	Name	SBO
S15	@nuc::rb(S788P̃,S800Ũ,b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{34} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_34\_0}([S16], \text{vol}(\text{cell}), \text{kdeg\_e2fbound}) \quad (232)$$

$$\text{Function\_for\_reaction\_34\_0}([S16], \text{vol}(\text{cell}), \text{kdeg\_e2fbound}) = \frac{\text{kdeg\_e2fbound} \cdot [S16]}{\text{vol}(\text{cell})} \quad (233)$$

$$\text{Function\_for\_reaction\_34\_0}([S16], \text{vol}(\text{cell}), \text{kdeg\_e2fbound}) = \frac{\text{kdeg\_e2fbound} \cdot [S16]}{\text{vol}(\text{cell})} \quad (234)$$

### 8.35 Reaction `reaction_35`

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

**Name** `reaction_35`

## Reaction equation



## Reactants

Table 91: Properties of each reactant.

Id	Name	SBO
S1	@nuc::rb(S788Ũ,S800Ũ,b)	
S14	@nuc::e2f(b)	

## Modifiers

Table 92: Properties of each modifier.

Id	Name	SBO
S1	@nuc::rb(S788Ũ,S800Ũ,b)	
S14	@nuc::e2f(b)	

## Product

Table 93: Properties of each product.

Id	Name	SBO
S2	@nuc::e2f(b!1).rb(S788Ũ,S800Ũ,b!1)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{35} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_35\_0}([S1], [S14], \text{vol}(\text{cell}), \text{kb\_rbe2f}) \quad (236)$$

$$\text{Function\_for\_reaction\_35\_0}([S1], [S14], \text{vol}(\text{cell}), \text{kb\_rbe2f}) = \frac{\text{kb\_rbe2f} \cdot [S1] \cdot [S14]}{\text{vol}(\text{cell})} \quad (237)$$

$$\text{Function\_for\_reaction\_35\_0}([S1], [S14], \text{vol}(\text{cell}), \text{kb\_rbe2f}) = \frac{\text{kb\_rbe2f} \cdot [S1] \cdot [S14]}{\text{vol}(\text{cell})} \quad (238)$$

### 8.36 Reaction reaction\_36

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

**Name** reaction\_36

#### Reaction equation



#### Reactants

Table 94: Properties of each reactant.

Id	Name	SBO
S14	@nuc::e2f(b)	
S15	@nuc::rb(S788P̃,S800Ũ,b)	

#### Modifiers

Table 95: Properties of each modifier.

Id	Name	SBO
S14	@nuc::e2f(b)	
S15	@nuc::rb(S788P̃,S800Ũ,b)	

#### Product

Table 96: Properties of each product.

Id	Name	SBO
S16	@nuc::e2f(b!1).rb(S788P̃,S800Ũ,b!1)	

#### Kinetic Law

**Derived unit** contains undeclared units

$$v_{36} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_36\_0}([S14], [S15], \text{vol}(\text{cell}), \text{kb\_rbpe2f}) \quad (240)$$

$$\text{Function\_for\_reaction\_36\_0}([S14], [S15], \text{vol}(\text{cell}), \text{kb\_rbpe2f}) = \frac{\text{kb\_rbpe2f} \cdot [S14] \cdot [S15]}{\text{vol}(\text{cell})} \quad (241)$$

$$\text{Function\_for\_reaction\_36\_0}([S14], [S15], \text{vol}(\text{cell}), \text{kb\_rbpe2f}) = \frac{\text{kb\_rbpe2f} \cdot [S14] \cdot [S15]}{\text{vol}(\text{cell})} \quad (242)$$

### 8.37 Reaction `reaction_37`

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

**Name** `reaction_37`

#### Reaction equation



#### Reactant

Table 97: Properties of each reactant.

Id	Name	SBO
S16	@nuc::e2f(b!1).rb(S788P̃,S800Ũ,b!1)	

#### Modifier

Table 98: Properties of each modifier.

Id	Name	SBO
S16	@nuc::e2f(b!1).rb(S788P̃,S800Ũ,b!1)	

#### Products

Table 99: Properties of each product.

Id	Name	SBO
S14	@nuc::e2f(b)	
S15	@nuc::rb(S788P̃,S800Ũ,b)	

#### Kinetic Law

**Derived unit** contains undeclared units

$$v_{37} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_37\_0}([S16], \text{vol}(\text{cell}), \text{kd\_rbpe2f}) \quad (244)$$

$$\text{Function\_for\_reaction\_37\_0}([S16], \text{vol}(\text{cell}), \text{kd\_rbpe2f}) = \frac{\text{kd\_rbpe2f} \cdot [S16]}{\text{vol}(\text{cell})} \quad (245)$$

$$\text{Function\_for\_reaction\_37\_0}([S16], \text{vol}(\text{cell}), \text{kd\_rbpe2f}) = \frac{\text{kd\_rbpe2f} \cdot [S16]}{\text{vol}(\text{cell})} \quad (246)$$

### 8.38 Reaction `reaction_38`

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

**Name** `reaction_38`

#### Reaction equation



#### Reactant

Table 100: Properties of each reactant.

Id	Name	SBO
S15	@nuc::rb(S788P̃,S800Ũ,b)	

#### Modifiers

Table 101: Properties of each modifier.

Id	Name	SBO
S18	@nuc::C2E(T160P̃,b)	
S15	@nuc::rb(S788P̃,S800Ũ,b)	

#### Product

Table 102: Properties of each product.

Id	Name	SBO
S21	@nuc::rb(S788P̃,S800P̃,b)	

#### Kinetic Law

**Derived unit** contains undeclared units

$$v_{38} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_38\_0}(\text{Km\_prb}, [\text{S15}], [\text{S18}], \text{vol}(\text{cell}), \text{kcatp\_rbc2}, \text{nrb}) \quad (248)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_38\_0}(\text{Km\_prb}, [\text{S15}], [\text{S18}], \text{vol}(\text{cell}), \text{kcatp\_rbc2}, \text{nrb}) \\ &= \frac{\frac{\text{kcatp\_rbc2} \cdot [\text{S18}] \cdot [\text{S15}]^{\text{nrb}}}{\text{Km\_prb}^{\text{nrb}} + [\text{S15}]^{\text{nrb}}}}{\text{vol}(\text{cell})} \end{aligned} \quad (249)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_38\_0}(\text{Km\_prb}, [\text{S15}], [\text{S18}], \text{vol}(\text{cell}), \text{kcatp\_rbc2}, \text{nrb}) \\ &= \frac{\frac{\text{kcatp\_rbc2} \cdot [\text{S18}] \cdot [\text{S15}]^{\text{nrb}}}{\text{Km\_prb}^{\text{nrb}} + [\text{S15}]^{\text{nrb}}}}{\text{vol}(\text{cell})} \end{aligned} \quad (250)$$

### 8.39 Reaction `reaction_39`

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

**Name** `reaction_39`

#### Reaction equation



#### Reactant

Table 103: Properties of each reactant.

Id	Name	SBO
S16	@nuc::e2f(b!1).rb(S788P̄,S800Ũ,b!1)	

#### Modifiers

Table 104: Properties of each modifier.

Id	Name	SBO
S18	@nuc::C2E(T160P̄,b)	
S16	@nuc::e2f(b!1).rb(S788P̄,S800Ũ,b!1)	

#### Products

Table 105: Properties of each product.

Id	Name	SBO
S14	@nuc::e2f(b)	
S21	@nuc::rb(S788P̃,S800P̃,b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{39} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_39\_0}(\text{Km\_prb}, [\text{S16}], [\text{S18}], \text{vol}(\text{cell}), \text{kcatp\_rbc2}, \text{nrb}) \quad (252)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_39\_0}(\text{Km\_prb}, [\text{S16}], [\text{S18}], \text{vol}(\text{cell}), \text{kcatp\_rbc2}, \text{nrb}) \\ &= \frac{\frac{\text{kcatp\_rbc2} \cdot [\text{S18}] \cdot [\text{S16}]^{\text{nrb}}}{\text{Km\_prb}^{\text{nrb}} + [\text{S16}]^{\text{nrb}}}}{\text{vol}(\text{cell})} \end{aligned} \quad (253)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_39\_0}(\text{Km\_prb}, [\text{S16}], [\text{S18}], \text{vol}(\text{cell}), \text{kcatp\_rbc2}, \text{nrb}) \\ &= \frac{\frac{\text{kcatp\_rbc2} \cdot [\text{S18}] \cdot [\text{S16}]^{\text{nrb}}}{\text{Km\_prb}^{\text{nrb}} + [\text{S16}]^{\text{nrb}}}}{\text{vol}(\text{cell})} \end{aligned} \quad (254)$$

## 8.40 Reaction reaction\_40

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

**Name** reaction\_40

## Reaction equation



## Reactant

Table 106: Properties of each reactant.

Id	Name	SBO
S15	@nuc::rb(S788P̃,S800P̃,b)	

## Modifiers



Table 107: Properties of each modifier.

Id	Name	SBO
S18	@nuc::C2E(T160P̃,b)	
S15	@nuc::rb(S788P̃,S800Ũ,b)	

## Product

Table 108: Properties of each product.

Id	Name	SBO
S1	@nuc::rb(S788Ũ,S800Ũ,b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{40} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_40\_0}(\text{Km\_dprb}, [\text{S15}], [\text{S18}], \text{vol}(\text{cell}), \text{kcatdp\_rbc4}, \text{kinh\_pp1}, \text{nrb}) \quad (256)$$

$$\begin{aligned} &\text{Function\_for\_reaction\_40\_0}(\text{Km\_dprb}, [\text{S15}], [\text{S18}], \\ &\quad \text{vol}(\text{cell}), \text{kcatdp\_rbc4}, \text{kinh\_pp1}, \text{nrb}) = \frac{\frac{\text{kcatdp\_rbc4} \cdot [\text{S15}]^{\text{nrb}}}{\text{Km\_dprb}^{\text{nrb}} + [\text{S15}]^{\text{nrb}}} \cdot 1}{1 + \text{kinh\_pp1} \cdot [\text{S18}]} \cdot \text{vol}(\text{cell}) \end{aligned} \quad (257)$$

$$\begin{aligned} &\text{Function\_for\_reaction\_40\_0}(\text{Km\_dprb}, [\text{S15}], [\text{S18}], \\ &\quad \text{vol}(\text{cell}), \text{kcatdp\_rbc4}, \text{kinh\_pp1}, \text{nrb}) = \frac{\frac{\text{kcatdp\_rbc4} \cdot [\text{S15}]^{\text{nrb}}}{\text{Km\_dprb}^{\text{nrb}} + [\text{S15}]^{\text{nrb}}} \cdot 1}{1 + \text{kinh\_pp1} \cdot [\text{S18}]} \cdot \text{vol}(\text{cell}) \end{aligned} \quad (258)$$

### 8.41 Reaction `reaction_41`

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

**Name** `reaction_41`

#### Reaction equation



## Reactant

Table 109: Properties of each reactant.

Id	Name	SBO
S16	@nuc::e2f(b!1).rb(S788P̃,S800Ũ,b!1)	

## Modifiers

Table 110: Properties of each modifier.

Id	Name	SBO
S18	@nuc::C2E(T160P̃,b)	
S16	@nuc::e2f(b!1).rb(S788P̃,S800Ũ,b!1)	

## Product

Table 111: Properties of each product.

Id	Name	SBO
S2	@nuc::e2f(b!1).rb(S788Ũ,S800Ũ,b!1)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{41} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_41\_0}(\text{Km\_dprb}, [\text{S16}], [\text{S18}], \text{vol}(\text{cell}), \text{kcatdp\_rbc4}, \text{kinh\_pp1}, \text{nrb}) \quad (260)$$

$$\begin{aligned} &\text{Function\_for\_reaction\_41\_0}(\text{Km\_dprb}, [\text{S16}], [\text{S18}], \\ &\text{vol}(\text{cell}), \text{kcatdp\_rbc4}, \text{kinh\_pp1}, \text{nrb}) = \frac{\frac{\text{kcatdp\_rbc4} \cdot [\text{S16}]^{\text{nrb}}}{\text{Km\_dprb}^{\text{nrb}} + [\text{S16}]^{\text{nrb}}} \cdot 1}{1 + \text{kinh\_pp1} \cdot [\text{S18}]} \cdot \text{vol}(\text{cell}) \end{aligned} \quad (261)$$

$$\begin{aligned} &\text{Function\_for\_reaction\_41\_0}(\text{Km\_dprb}, [\text{S16}], [\text{S18}], \\ &\text{vol}(\text{cell}), \text{kcatdp\_rbc4}, \text{kinh\_pp1}, \text{nrb}) = \frac{\frac{\text{kcatdp\_rbc4} \cdot [\text{S16}]^{\text{nrb}}}{\text{Km\_dprb}^{\text{nrb}} + [\text{S16}]^{\text{nrb}}} \cdot 1}{1 + \text{kinh\_pp1} \cdot [\text{S18}]} \cdot \text{vol}(\text{cell}) \end{aligned} \quad (262)$$

### 8.42 Reaction [reaction\\_42](#)

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

**Name** reaction\_42

## Reaction equation



## Reactant

Table 112: Properties of each reactant.

Id	Name	SBO
S17	@nuc::dnapre1()	

## Modifier

Table 113: Properties of each modifier.

Id	Name	SBO
S17	@nuc::dnapre1()	

## Product

Table 114: Properties of each product.

Id	Name	SBO
S22	@nuc::dnapre2()	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{42} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_42\_0}([S17], \text{vol}(\text{cell}), k\_delay) \quad (264)$$

$$\text{Function\_for\_reaction\_42\_0}([S17], \text{vol}(\text{cell}), k\_delay) = \frac{k\_delay \cdot [S17]}{\text{vol}(\text{cell})} \quad (265)$$

$$\text{Function\_for\_reaction\_42\_0}([S17], \text{vol}(\text{cell}), k\_delay) = \frac{k\_delay \cdot [S17]}{\text{vol}(\text{cell})} \quad (266)$$

### 8.43 Reaction `reaction_43`

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

**Name** `reaction_43`

## Reaction equation



## Reactant

Table 115: Properties of each reactant.

Id	Name	SBO
S19	@cyto::C4D1(b!1).p21(b!1)	

## Modifier

Table 116: Properties of each modifier.

Id	Name	SBO
S19	@cyto::C4D1(b!1).p21(b!1)	

## Product

Table 117: Properties of each product.

Id	Name	SBO
S12	@cyto::p21(b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{43} = \text{vol}(\text{cell}) \cdot \text{Function\_for\_reaction\_43\_0}([S19], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c4}, \text{kdeg\_c4gsk3b}) \quad (268)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_43\_0}([S19], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c4}, \text{kdeg\_c4gsk3b}) \\ &= \frac{(\text{kdeg\_c4} + \text{kdeg\_c4gsk3b} \cdot \text{gsk3b}) \cdot [S19]}{\text{vol}(\text{cell})} \end{aligned} \quad (269)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_43\_0}([S19], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c4}, \text{kdeg\_c4gsk3b}) \\ &= \frac{(\text{kdeg\_c4} + \text{kdeg\_c4gsk3b} \cdot \text{gsk3b}) \cdot [S19]}{\text{vol}(\text{cell})} \end{aligned} \quad (270)$$

## 8.44 Reaction `reaction_44`

This is an irreversible reaction of one reactant forming no product influenced by one modifier.

**Name** `reaction_44`

### Reaction equation



### Reactant

Table 118: Properties of each reactant.

Id	Name	SBO
S18	@nuc::C2E(T160P̃,b)	

### Modifier

Table 119: Properties of each modifier.

Id	Name	SBO
S18	@nuc::C2E(T160P̃,b)	

### Kinetic Law

**Derived unit** contains undeclared units

$$\begin{aligned} v_{44} &= \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_44\_0}([S18], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \\ &\quad (272) \end{aligned}$$

$$\begin{aligned} &\text{Function\_for\_reaction\_44\_0}([S18], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \\ &= \frac{(\text{kdeg\_c2} + \text{kdeg\_c2gsk3b} \cdot \text{gsk3b}) \cdot [S18]}{\text{vol}(\text{cell})} \quad (273) \end{aligned}$$

$$\begin{aligned} &\text{Function\_for\_reaction\_44\_0}([S18], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \\ &= \frac{(\text{kdeg\_c2} + \text{kdeg\_c2gsk3b} \cdot \text{gsk3b}) \cdot [S18]}{\text{vol}(\text{cell})} \quad (274) \end{aligned}$$

## 8.45 Reaction `reaction_45`

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

**Name** `reaction_45`

### Reaction equation



### Reactant

Table 120: Properties of each reactant.

Id	Name	SBO
S20	@cyto::C2E(T160Û,b!1).p21(b!1)	

### Modifier

Table 121: Properties of each modifier.

Id	Name	SBO
S20	@cyto::C2E(T160Û,b!1).p21(b!1)	

### Product

Table 122: Properties of each product.

Id	Name	SBO
S12	@cyto::p21(b)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{45} = \text{vol}(\text{cell}) \cdot \text{Function\_for\_reaction\_45\_0}([S20], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \quad (276)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_45\_0}([S20], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \\ &= \frac{(\text{kdeg\_c2} + \text{kdeg\_c2gsk3b} \cdot \text{gsk3b}) \cdot [S20]}{\text{vol}(\text{cell})} \end{aligned} \quad (277)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_45\_0}([S20], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \\ &= \frac{(\text{kdeg\_c2} + \text{kdeg\_c2gsk3b} \cdot \text{gsk3b}) \cdot [S20]}{\text{vol}(\text{cell})} \end{aligned} \quad (278)$$

## 8.46 Reaction `reaction_46`

This is an irreversible reaction of one reactant forming no product influenced by one modifier.

**Name** `reaction_46`

### Reaction equation



### Reactant

Table 123: Properties of each reactant.

Id	Name	SBO
S18	@nuc::C2E(T160P̃,b)	

### Modifier

Table 124: Properties of each modifier.

Id	Name	SBO
S18	@nuc::C2E(T160P̃,b)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{46} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_46\_0}([S18], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2c2gsk3b}) \quad (280)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_46\_0}([S18], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2c2gsk3b}) \\ &= \frac{\text{kdeg\_c2c2gsk3b} \cdot \text{gsk3b} \cdot [S18]}{\text{vol}(\text{cell})} \end{aligned} \quad (281)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_46\_0}([S18], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2c2gsk3b}) \\ &= \frac{\text{kdeg\_c2c2gsk3b} \cdot \text{gsk3b} \cdot [S18]}{\text{vol}(\text{cell})} \end{aligned} \quad (282)$$

### 8.47 Reaction `reaction_47`

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

**Name** `reaction_47`

#### Reaction equation



#### Reactant

Table 125: Properties of each reactant.

Id	Name	SBO
S18	@nuc::C2E(T160P̃,b)	

#### Modifier

Table 126: Properties of each modifier.

Id	Name	SBO
S18	@nuc::C2E(T160P̃,b)	

#### Product

Table 127: Properties of each product.

Id	Name	SBO
S13	@nuc::C2E(T160Ũ,b)	

#### Kinetic Law

**Derived unit** contains undeclared units

$$v_{47} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_47\_0}([\text{S18}], \text{vol}(\text{cell}), \text{kdp\_c2cak}) \quad (284)$$

$$\text{Function\_for\_reaction\_47\_0}([\text{S18}], \text{vol}(\text{cell}), \text{kdp\_c2cak}) = \frac{\text{kdp\_c2cak} \cdot [\text{S18}]}{\text{vol}(\text{cell})} \quad (285)$$

$$\text{Function\_for\_reaction\_47\_0}([\text{S18}], \text{vol}(\text{cell}), \text{kdp\_c2cak}) = \frac{\text{kdp\_c2cak} \cdot [\text{S18}]}{\text{vol}(\text{cell})} \quad (286)$$



## 8.48 Reaction `reaction_48`

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

**Name** `reaction_48`

### Reaction equation



### Reactant

Table 128: Properties of each reactant.

Id	Name	SBO
S19	@cyto::C4D1(b!1).p21(b!1)	

### Modifier

Table 129: Properties of each modifier.

Id	Name	SBO
S19	@cyto::C4D1(b!1).p21(b!1)	

### Products

Table 130: Properties of each product.

Id	Name	SBO
S10	@cyto::C4D1(b)	
S12	@cyto::p21(b)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{48} = \text{vol}(\text{cell}) \cdot \text{Function\_for\_reaction\_48\_0}([S19], \text{vol}(\text{cell}), \text{kd\_p21c4}) \quad (288)$$

$$\text{Function\_for\_reaction\_48\_0}([S19], \text{vol}(\text{cell}), \text{kd\_p21c4}) = \frac{\text{kd\_p21c4} \cdot [S19]}{\text{vol}(\text{cell})} \quad (289)$$

$$\text{Function\_for\_reaction\_48\_0}([S19], \text{vol}(\text{cell}), \text{kd\_p21c4}) = \frac{\text{kd\_p21c4} \cdot [S19]}{\text{vol}(\text{cell})} \quad (290)$$

## 8.49 Reaction reaction\_49

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

**Name** reaction\_49

### Reaction equation



### Reactants

Table 131: Properties of each reactant.

Id	Name	SBO
S11	@nuc::p21(b)	
S18	@nuc::C2E(T160P̃,b)	

### Modifiers

Table 132: Properties of each modifier.

Id	Name	SBO
S11	@nuc::p21(b)	
S18	@nuc::C2E(T160P̃,b)	

### Product

Table 133: Properties of each product.

Id	Name	SBO
S23	@nuc::C2E(T160P̃,b!1),p21(b!1)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{49} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_49\_0}([S11], [S18], \text{vol}(\text{cell}), \text{kb\_p21c2}) \quad (292)$$

$$\text{Function\_for\_reaction\_49\_0}([S11], [S18], \text{vol}(\text{cell}), \text{kb\_p21c2}) = \frac{\text{kb\_p21c2} \cdot [S11] \cdot [S18]}{\text{vol}(\text{cell})} \quad (293)$$

$$\text{Function\_for\_reaction\_49\_0}([S11],[S18],\text{vol}(\text{cell}),\text{kb\_p21c2}) = \frac{\text{kb\_p21c2} \cdot [S11] \cdot [S18]}{\text{vol}(\text{cell})} \quad (294)$$

## 8.50 Reaction `reaction_50`

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

**Name** `reaction_50`

### Reaction equation



### Reactant

Table 134: Properties of each reactant.

Id	Name	SBO
S20	@cyto::C2E(T160Û,b!1).p21(b!1)	

### Modifier

Table 135: Properties of each modifier.

Id	Name	SBO
S20	@cyto::C2E(T160Û,b!1).p21(b!1)	

### Products

Table 136: Properties of each product.

Id	Name	SBO
S4	@cyto::C2E(T160Û,b)	
S12	@cyto::p21(b)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{50} = \text{vol}(\text{cell}) \cdot \text{Function\_for\_reaction\_50\_0}([S20],\text{vol}(\text{cell}),\text{kd\_p21c2}) \quad (296)$$

$$\text{Function\_for\_reaction\_50\_0}([S20], \text{vol}(\text{cell}), \text{kd\_p21c2}) = \frac{\text{kd\_p21c2} \cdot [S20]}{\text{vol}(\text{cell})} \quad (297)$$

$$\text{Function\_for\_reaction\_50\_0}([S20], \text{vol}(\text{cell}), \text{kd\_p21c2}) = \frac{\text{kd\_p21c2} \cdot [S20]}{\text{vol}(\text{cell})} \quad (298)$$

### 8.51 Reaction `reaction_51`

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

**Name** `reaction_51`

#### Reaction equation



#### Reactant

Table 137: Properties of each reactant.

Id	Name	SBO
S19	@cyto::C4D1(b!1).p21(b!1)	

#### Modifier

Table 138: Properties of each modifier.

Id	Name	SBO
S19	@cyto::C4D1(b!1).p21(b!1)	

#### Product

Table 139: Properties of each product.

Id	Name	SBO
S24	@nuc::C4D1(b!1).p21(b!1)	

#### Kinetic Law

**Derived unit** contains undeclared units

$$v_{51} = \text{Function\_for\_reaction\_51\_0}([S19], \text{Vratio}, \text{vol}(\text{cell}), \text{kimport}) \quad (300)$$

$$\text{Function\_for\_reaction\_51\_0}([S19], V_{\text{ratio}}, \text{vol}(\text{cell}), k_{\text{import}}) = \frac{\frac{k_{\text{import}}}{V_{\text{ratio}}} \cdot [S19]}{\text{vol}(\text{cell})} \quad (301)$$

## 8.52 Reaction `reaction_52`

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

**Name** `reaction_52`

### Reaction equation



### Reactant

Table 140: Properties of each reactant.

Id	Name	SBO
S20	@cyto::C2E(T160Û,b!1).p21(b!1)	

### Modifier

Table 141: Properties of each modifier.

Id	Name	SBO
S20	@cyto::C2E(T160Û,b!1).p21(b!1)	

### Product

Table 142: Properties of each product.

Id	Name	SBO
S3	@nuc::C2E(T160Û,b!1).p21(b!1)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{52} = \text{Function\_for\_reaction\_52\_0}([S20], V_{\text{ratio}}, \text{vol}(\text{cell}), k_{\text{import}}) \quad (303)$$

$$\text{Function\_for\_reaction\_52\_0}([S20], V_{\text{ratio}}, \text{vol}(\text{cell}), k_{\text{import}}) = \frac{\frac{k_{\text{import}}}{V_{\text{ratio}}} \cdot [S20]}{\text{vol}(\text{cell})} \quad (304)$$

### 8.53 Reaction `reaction_53`

This is a reversible reaction of one reactant forming no product influenced by one modifier.

**Name** `reaction_53`

#### Reaction equation



#### Reactant

Table 143: Properties of each reactant.

Id	Name	SBO
S19	@cyto::C4D1(b!1).p21(b!1)	

#### Modifier

Table 144: Properties of each modifier.

Id	Name	SBO
S19	@cyto::C4D1(b!1).p21(b!1)	

#### Kinetic Law

**Derived unit** contains undeclared units

$$v_{53} = \text{vol}(\text{cell}) \cdot \text{Function\_for\_reaction\_53\_0}([S19], V_{\text{ratio}}, \text{vol}(\text{cell}), \text{kimport}) \quad (306)$$

$$\text{Function\_for\_reaction\_53\_0}([S19], V_{\text{ratio}}, \text{vol}(\text{cell}), \text{kimport}) = \frac{\text{kimport} \cdot \left(1 - \frac{1}{V_{\text{ratio}}}\right) \cdot [S19]}{\text{vol}(\text{cell})} \quad (307)$$

$$\text{Function\_for\_reaction\_53\_0}([S19], V_{\text{ratio}}, \text{vol}(\text{cell}), \text{kimport}) = \frac{\text{kimport} \cdot \left(1 - \frac{1}{V_{\text{ratio}}}\right) \cdot [S19]}{\text{vol}(\text{cell})} \quad (308)$$

### 8.54 Reaction `reaction_54`

This is a reversible reaction of one reactant forming no product influenced by one modifier.

**Name** `reaction_54`

### Reaction equation



### Reactant

Table 145: Properties of each reactant.

Id	Name	SBO
S20	@cyto::C2E(T160Û,b!1).p21(b!1)	

### Modifier

Table 146: Properties of each modifier.

Id	Name	SBO
S20	@cyto::C2E(T160Û,b!1).p21(b!1)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{54} = \text{vol}(\text{cell}) \cdot \text{Function\_for\_reaction\_54\_0}([S20], V_{\text{ratio}}, \text{vol}(\text{cell}), \text{kimport}) \quad (310)$$

$$\text{Function\_for\_reaction\_54\_0}([S20], V_{\text{ratio}}, \text{vol}(\text{cell}), \text{kimport}) = \frac{\text{kimport} \cdot \left(1 - \frac{1}{V_{\text{ratio}}}\right) \cdot [S20]}{\text{vol}(\text{cell})} \quad (311)$$

$$\text{Function\_for\_reaction\_54\_0}([S20], V_{\text{ratio}}, \text{vol}(\text{cell}), \text{kimport}) = \frac{\text{kimport} \cdot \left(1 - \frac{1}{V_{\text{ratio}}}\right) \cdot [S20]}{\text{vol}(\text{cell})} \quad (312)$$

### 8.55 Reaction [reaction\\_55](#)

This is an irreversible reaction of one reactant forming no product influenced by one modifier.

**Name** [reaction\\_55](#)

### Reaction equation



### Reactant

Table 147: Properties of each reactant.

Id	Name	SBO
S21	@nuc::rb(S788P̃,S800P̃,b)	

## Modifier

Table 148: Properties of each modifier.

Id	Name	SBO
S21	@nuc::rb(S788P̃,S800P̃,b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{55} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_55\_0}([S21], \text{vol}(\text{cell}), \text{kdeg\_rbfree}) \quad (314)$$

$$\text{Function\_for\_reaction\_55\_0}([S21], \text{vol}(\text{cell}), \text{kdeg\_rbfree}) = \frac{\text{kdeg\_rbfree} \cdot [S21]}{\text{vol}(\text{cell})} \quad (315)$$

$$\text{Function\_for\_reaction\_55\_0}([S21], \text{vol}(\text{cell}), \text{kdeg\_rbfree}) = \frac{\text{kdeg\_rbfree} \cdot [S21]}{\text{vol}(\text{cell})} \quad (316)$$

## 8.56 Reaction `reaction_56`

This is an irreversible reaction of one reactant forming no product influenced by two modifiers.

**Name** `reaction_56`

## Reaction equation



## Reactant

Table 149: Properties of each reactant.

Id	Name	SBO
S21	@nuc::rb(S788P̃,S800P̃,b)	



## Modifiers

Table 150: Properties of each modifier.

Id	Name	SBO
S11	@nuc::p21(b)	
S21	@nuc::rb(S788P̃,S800P̃,b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{56} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_56\_0}([S11], [S21], \text{vol}(\text{cell}), \text{kdeg\_rbp21}) \quad (318)$$

$$\text{Function\_for\_reaction\_56\_0}([S11], [S21], \text{vol}(\text{cell}), \text{kdeg\_rbp21}) = \frac{\text{kdeg\_rbp21} \cdot [S11] \cdot [S21]}{\text{vol}(\text{cell})} \quad (319)$$

$$\text{Function\_for\_reaction\_56\_0}([S11], [S21], \text{vol}(\text{cell}), \text{kdeg\_rbp21}) = \frac{\text{kdeg\_rbp21} \cdot [S11] \cdot [S21]}{\text{vol}(\text{cell})} \quad (320)$$

### 8.57 Reaction `reaction_57`

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

**Name** `reaction_57`

#### Reaction equation



## Reactant

Table 151: Properties of each reactant.

Id	Name	SBO
S21	@nuc::rb(S788P̃,S800P̃,b)	

## Modifiers

Table 152: Properties of each modifier.

Id	Name	SBO
S18	@nuc::C2E(T160P̃,b)	
S21	@nuc::rb(S788P̃,S800P̃,b)	

## Product

Table 153: Properties of each product.

Id	Name	SBO
S15	@nuc::rb(S788P̃,S800P̃,b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{57} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_57\_0}(\text{Km\_dprb}, [\text{S18}], [\text{S21}], \text{vol}(\text{cell}), \text{kcatdp\_rbc2}, \text{kinh\_pp1}, \text{nrb}) \quad (322)$$

$$\begin{aligned} &\text{Function\_for\_reaction\_57\_0}(\text{Km\_dprb}, [\text{S18}], [\text{S21}], \\ &\text{vol}(\text{cell}), \text{kcatdp\_rbc2}, \text{kinh\_pp1}, \text{nrb}) = \frac{\frac{\text{kcatdp\_rbc2} \cdot [\text{S21}]^{\text{nrb}}}{\text{Km\_dprb}^{\text{nrb}} + [\text{S21}]^{\text{nrb}}} \cdot 1}{1 + \text{kinh\_pp1} \cdot [\text{S18}]} \cdot \text{vol}(\text{cell}) \end{aligned} \quad (323)$$

$$\begin{aligned} &\text{Function\_for\_reaction\_57\_0}(\text{Km\_dprb}, [\text{S18}], [\text{S21}], \\ &\text{vol}(\text{cell}), \text{kcatdp\_rbc2}, \text{kinh\_pp1}, \text{nrb}) = \frac{\frac{\text{kcatdp\_rbc2} \cdot [\text{S21}]^{\text{nrb}}}{\text{Km\_dprb}^{\text{nrb}} + [\text{S21}]^{\text{nrb}}} \cdot 1}{1 + \text{kinh\_pp1} \cdot [\text{S18}]} \cdot \text{vol}(\text{cell}) \end{aligned} \quad (324)$$

## 8.58 Reaction [reaction\\_58](#)

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

**Name** reaction\_58

## Reaction equation



## Reactant

Table 154: Properties of each reactant.

Id	Name	SBO
S22	@nuc::dnapre2()	

### Modifier

Table 155: Properties of each modifier.

Id	Name	SBO
S22	@nuc::dnapre2()	

### Product

Table 156: Properties of each product.

Id	Name	SBO
S25	@nuc::dnapre3()	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{58} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_58\_0}([S22], \text{vol}(\text{cell}), k_{\text{delay}}) \quad (326)$$

$$\text{Function\_for\_reaction\_58\_0}([S22], \text{vol}(\text{cell}), k_{\text{delay}}) = \frac{k_{\text{delay}} \cdot [S22]}{\text{vol}(\text{cell})} \quad (327)$$

$$\text{Function\_for\_reaction\_58\_0}([S22], \text{vol}(\text{cell}), k_{\text{delay}}) = \frac{k_{\text{delay}} \cdot [S22]}{\text{vol}(\text{cell})} \quad (328)$$

## 8.59 Reaction `reaction_59`

This is an irreversible reaction of one reactant forming no product influenced by one modifier.

**Name** `reaction_59`

### Reaction equation



### Reactant

Table 157: Properties of each reactant.

Id	Name	SBO
S24	@nuc::C4D1(b!1).p21(b!1)	

### Modifier

Table 158: Properties of each modifier.

Id	Name	SBO
S24	@nuc::C4D1(b!1).p21(b!1)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{59} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_59\_0}([S24], \text{vol}(\text{cell}), \text{kdeg\_c4}) \quad (330)$$

$$\text{Function\_for\_reaction\_59\_0}([S24], \text{vol}(\text{cell}), \text{kdeg\_c4}) = \frac{\text{kdeg\_c4} \cdot [S24]}{\text{vol}(\text{cell})} \quad (331)$$

$$\text{Function\_for\_reaction\_59\_0}([S24], \text{vol}(\text{cell}), \text{kdeg\_c4}) = \frac{\text{kdeg\_c4} \cdot [S24]}{\text{vol}(\text{cell})} \quad (332)$$

## 8.60 Reaction `reaction_60`

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

**Name** `reaction_60`

### Reaction equation



### Reactant

Table 159: Properties of each reactant.

Id	Name	SBO
S23	@nuc::C2E(T160P̄,b!1).p21(b!1)	

## Modifier

Table 160: Properties of each modifier.

Id	Name	SBO
S23	@nuc::C2E(T160P̃,b!1),p21(b!1)	

## Product

Table 161: Properties of each product.

Id	Name	SBO
S11	@nuc::p21(b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{60} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_60\_0}([S23], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \quad (334)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_60\_0}([S23], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \\ &= \frac{(\text{kdeg\_c2} + \text{kdeg\_c2gsk3b} \cdot \text{gsk3b}) \cdot [S23]}{\text{vol}(\text{cell})} \end{aligned} \quad (335)$$

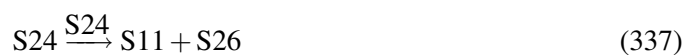
$$\begin{aligned} & \text{Function\_for\_reaction\_60\_0}([S23], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c2}, \text{kdeg\_c2gsk3b}) \\ &= \frac{(\text{kdeg\_c2} + \text{kdeg\_c2gsk3b} \cdot \text{gsk3b}) \cdot [S23]}{\text{vol}(\text{cell})} \end{aligned} \quad (336)$$

### 8.61 Reaction `reaction_61`

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

**Name** `reaction_61`

#### Reaction equation



## Reactant

Table 162: Properties of each reactant.

Id	Name	SBO
S24	@nuc::C4D1(b!1).p21(b!1)	

## Modifier

Table 163: Properties of each modifier.

Id	Name	SBO
S24	@nuc::C4D1(b!1).p21(b!1)	

## Products

Table 164: Properties of each product.

Id	Name	SBO
S11	@nuc::p21(b)	
S26	@nuc::C4D1(b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{61} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_61\_0}([S24], \text{vol}(\text{cell}), \text{kd\_p21c4}) \quad (338)$$

$$\text{Function\_for\_reaction\_61\_0}([S24], \text{vol}(\text{cell}), \text{kd\_p21c4}) = \frac{\text{kd\_p21c4} \cdot [S24]}{\text{vol}(\text{cell})} \quad (339)$$

$$\text{Function\_for\_reaction\_61\_0}([S24], \text{vol}(\text{cell}), \text{kd\_p21c4}) = \frac{\text{kd\_p21c4} \cdot [S24]}{\text{vol}(\text{cell})} \quad (340)$$

## 8.62 Reaction [reaction\\_62](#)

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

**Name** [reaction\\_62](#)

## Reaction equation



## Reactant

Table 165: Properties of each reactant.

Id	Name	SBO
S23	@nuc::C2E(T160P̃,b!1).p21(b!1)	

### Modifier

Table 166: Properties of each modifier.

Id	Name	SBO
S23	@nuc::C2E(T160P̃,b!1).p21(b!1)	

### Products

Table 167: Properties of each product.

Id	Name	SBO
S11	@nuc::p21(b)	
S18	@nuc::C2E(T160P̃,b)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{62} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_62\_0}([S23], \text{vol}(\text{cell}), \text{kd\_p21c2}) \quad (342)$$

$$\text{Function\_for\_reaction\_62\_0}([S23], \text{vol}(\text{cell}), \text{kd\_p21c2}) = \frac{\text{kd\_p21c2} \cdot [S23]}{\text{vol}(\text{cell})} \quad (343)$$

$$\text{Function\_for\_reaction\_62\_0}([S23], \text{vol}(\text{cell}), \text{kd\_p21c2}) = \frac{\text{kd\_p21c2} \cdot [S23]}{\text{vol}(\text{cell})} \quad (344)$$

### 8.63 Reaction `reaction_63`

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

**Name** `reaction_63`

### Reaction equation



### Reactant

Table 168: Properties of each reactant.

Id	Name	SBO
S23	@nuc::C2E(T160P̃,b!1),p21(b!1)	

## Modifiers

Table 169: Properties of each modifier.

Id	Name	SBO
S18	@nuc::C2E(T160P̃,b)	
S14	@nuc::e2f(b)	
S23	@nuc::C2E(T160P̃,b!1),p21(b!1)	

## Product

Table 170: Properties of each product.

Id	Name	SBO
S18	@nuc::C2E(T160P̃,b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{63} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_63\_0}([S14], [S18], [S23], \text{vol}(\text{cell}), \text{erk}, \text{kdeg\_p21c2skp2}, \text{kdeg\_p21erkskp2}, \text{kdeg\_p21skp2}) \quad (346)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_63\_0}([S14], [S18], [S23], \text{vol}(\text{cell}), \\ & \text{erk}, \text{kdeg\_p21c2skp2}, \text{kdeg\_p21erkskp2}, \text{kdeg\_p21skp2}) \\ &= \frac{(\text{kdeg\_p21erkskp2} \cdot \text{erk} + \text{kdeg\_p21c2skp2} \cdot [S18] + \text{kdeg\_p21skp2}) \cdot [S14] \cdot [S23]}{\text{vol}(\text{cell})} \end{aligned} \quad (347)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_63\_0}([S14], [S18], [S23], \text{vol}(\text{cell}), \\ & \text{erk}, \text{kdeg\_p21c2skp2}, \text{kdeg\_p21erkskp2}, \text{kdeg\_p21skp2}) \\ &= \frac{(\text{kdeg\_p21erkskp2} \cdot \text{erk} + \text{kdeg\_p21c2skp2} \cdot [S18] + \text{kdeg\_p21skp2}) \cdot [S14] \cdot [S23]}{\text{vol}(\text{cell})} \end{aligned} \quad (348)$$



## 8.64 Reaction `reaction_64`

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

**Name** `reaction_64`

### Reaction equation



### Reactant

Table 171: Properties of each reactant.

Id	Name	SBO
S24	@nuc::C4D1(b!1),p21(b!1)	

### Modifiers

Table 172: Properties of each modifier.

Id	Name	SBO
S18	@nuc::C2E(T160P̃,b)	
S14	@nuc::e2f(b)	
S24	@nuc::C4D1(b!1),p21(b!1)	

### Product

Table 173: Properties of each product.

Id	Name	SBO
S26	@nuc::C4D1(b)	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{64} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_64.0}([S14], [S18], [S24], \text{vol}(\text{cell}), \text{erk}, \text{kdeg\_p21c2skp2}, \text{kdeg\_p21erkskp2}, \text{kdeg\_p21skp2}) \quad (350)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_64\_0}([S14], [S18], [S24], \text{vol}(\text{cell}), \\ & \text{erk}, \text{kdeg\_p21c2skp2}, \text{kdeg\_p21erkskp2}, \text{kdeg\_p21skp2}) \\ &= \frac{(\text{kdeg\_p21erkskp2} \cdot \text{erk} + \text{kdeg\_p21c2skp2} \cdot [S18] + \text{kdeg\_p21skp2}) \cdot [S14] \cdot [S24]}{\text{vol}(\text{cell})} \end{aligned} \quad (351)$$

$$\begin{aligned} & \text{Function\_for\_reaction\_64\_0}([S14], [S18], [S24], \text{vol}(\text{cell}), \\ & \text{erk}, \text{kdeg\_p21c2skp2}, \text{kdeg\_p21erkskp2}, \text{kdeg\_p21skp2}) \\ &= \frac{(\text{kdeg\_p21erkskp2} \cdot \text{erk} + \text{kdeg\_p21c2skp2} \cdot [S18] + \text{kdeg\_p21skp2}) \cdot [S14] \cdot [S24]}{\text{vol}(\text{cell})} \end{aligned} \quad (352)$$

### 8.65 Reaction `reaction_65`

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

**Name** `reaction_65`

#### Reaction equation



#### Reactant

Table 174: Properties of each reactant.

Id	Name	SBO
S25	@nuc::dnapre3()	

#### Modifier

Table 175: Properties of each modifier.

Id	Name	SBO
S25	@nuc::dnapre3()	

#### Product

Table 176: Properties of each product.

Id	Name	SBO
S27	@nuc::dnapre4()	

### Kinetic Law

**Derived unit** contains undeclared units

$$v_{65} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_65\_0}([S25], \text{vol}(\text{cell}), k_{\text{delay}}) \quad (354)$$

$$\text{Function\_for\_reaction\_65\_0}([S25], \text{vol}(\text{cell}), k_{\text{delay}}) = \frac{k_{\text{delay}} \cdot [S25]}{\text{vol}(\text{cell})} \quad (355)$$

$$\text{Function\_for\_reaction\_65\_0}([S25], \text{vol}(\text{cell}), k_{\text{delay}}) = \frac{k_{\text{delay}} \cdot [S25]}{\text{vol}(\text{cell})} \quad (356)$$

### 8.66 Reaction reaction\_66

This is an irreversible reaction of one reactant forming no product influenced by one modifier.

**Name** reaction\_66

### Reaction equation



### Reactant

Table 177: Properties of each reactant.

Id	Name	SBO
S26	@nuc::C4D1(b)	

### Modifier

Table 178: Properties of each modifier.

Id	Name	SBO
S26	@nuc::C4D1(b)	

## Kinetic Law

**Derived unit** contains undeclared units

$$\begin{aligned} v_{66} &= \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_66\_0}([S26], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c4}, \text{kdeg\_c4gsk3b}) \\ &\quad (358) \end{aligned}$$

$$\begin{aligned} &\text{Function\_for\_reaction\_66\_0}([S26], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c4}, \text{kdeg\_c4gsk3b}) \\ &= \frac{(\text{kdeg\_c4} + \text{kdeg\_c4gsk3b} \cdot \text{gsk3b}) \cdot [S26]}{\text{vol}(\text{cell})} \end{aligned} \quad (359)$$

$$\begin{aligned} &\text{Function\_for\_reaction\_66\_0}([S26], \text{vol}(\text{cell}), \text{gsk3b}, \text{kdeg\_c4}, \text{kdeg\_c4gsk3b}) \\ &= \frac{(\text{kdeg\_c4} + \text{kdeg\_c4gsk3b} \cdot \text{gsk3b}) \cdot [S26]}{\text{vol}(\text{cell})} \end{aligned} \quad (360)$$

## 8.67 Reaction `reaction_67`

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

**Name** `reaction_67`

### Reaction equation



### Reactants

Table 179: Properties of each reactant.

Id	Name	SBO
S11	@nuc::p21(b)	
S26	@nuc::C4D1(b)	

### Modifiers

Table 180: Properties of each modifier.

Id	Name	SBO
S11	@nuc::p21(b)	
S26	@nuc::C4D1(b)	

## Product

Table 181: Properties of each product.

Id	Name	SBO
S24	@nuc::C4D1(b!1).p21(b!1)	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{67} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_67\_0}([S11], [S26], \text{vol}(\text{cell}), \text{kb\_p21c4}) \quad (362)$$

$$\text{Function\_for\_reaction\_67\_0}([S11], [S26], \text{vol}(\text{cell}), \text{kb\_p21c4}) = \frac{\text{kb\_p21c4} \cdot [S11] \cdot [S26]}{\text{vol}(\text{cell})} \quad (363)$$

$$\text{Function\_for\_reaction\_67\_0}([S11], [S26], \text{vol}(\text{cell}), \text{kb\_p21c4}) = \frac{\text{kb\_p21c4} \cdot [S11] \cdot [S26]}{\text{vol}(\text{cell})} \quad (364)$$

## 8.68 Reaction `reaction_68`

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

**Name** `reaction_68`

### Reaction equation



## Reactant

Table 182: Properties of each reactant.

Id	Name	SBO
S27	@nuc::dnapre4()	

## Modifier

Table 183: Properties of each modifier.

Id	Name	SBO
S27	@nuc::dnapre4()	

## Product

Table 184: Properties of each product.

Id	Name	SBO
S28	@nuc::dnapre5()	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{68} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_68\_0}([S27], \text{vol}(\text{cell}), k_{\text{delay}}) \quad (366)$$

$$\text{Function\_for\_reaction\_68\_0}([S27], \text{vol}(\text{cell}), k_{\text{delay}}) = \frac{k_{\text{delay}} \cdot [S27]}{\text{vol}(\text{cell})} \quad (367)$$

$$\text{Function\_for\_reaction\_68\_0}([S27], \text{vol}(\text{cell}), k_{\text{delay}}) = \frac{k_{\text{delay}} \cdot [S27]}{\text{vol}(\text{cell})} \quad (368)$$

## 8.69 Reaction `reaction_69`

This is an irreversible reaction of one reactant forming no product influenced by one modifier.

**Name** `reaction_69`

## Reaction equation



## Reactant

Table 185: Properties of each reactant.

Id	Name	SBO
S28	@nuc::dnapre5()	

## Modifier

Table 186: Properties of each modifier.

Id	Name	SBO
S28	@nuc::dnapre5()	

## Kinetic Law

**Derived unit** contains undeclared units

$$v_{69} = \text{vol}(\text{Nucleus}) \cdot \text{Function\_for\_reaction\_69\_0}([S28], \text{vol}(\text{cell}), k\_delay) \quad (370)$$

$$\text{Function\_for\_reaction\_69\_0}([S28], \text{vol}(\text{cell}), k\_delay) = \frac{k\_delay \cdot [S28]}{\text{vol}(\text{cell})} \quad (371)$$

$$\text{Function\_for\_reaction\_69\_0}([S28], \text{vol}(\text{cell}), k\_delay) = \frac{k\_delay \cdot [S28]}{\text{vol}(\text{cell})} \quad (372)$$

## 9 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.

### 9.1 Species S4

**Name** @cyto::C2E(T160Ū,b)

**Notes** Cyclin E:CDK2 complex - T160 phosphorylation

**Initial concentration** 0.415 nmol · l<sup>-1</sup>

This species takes part in six reactions (as a reactant in [reaction\\_4](#), [reaction\\_23](#) and as a product in [reaction\\_2](#), [reaction\\_50](#) and as a modifier in [reaction\\_4](#), [reaction\\_23](#)).

$$\frac{d}{dt}S4 = v_2 + v_{50} - v_4 - v_{23} \quad (373)$$

## 9.2 Species S10

**Name** @cyto::C4D1(b)

**Notes** Cyclin D1:CDK4 complex

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in six reactions (as a reactant in [reaction\\_19](#), [reaction\\_22](#) and as a product in [reaction\\_1](#), [reaction\\_48](#) and as a modifier in [reaction\\_19](#), [reaction\\_22](#)).

$$\frac{d}{dt}S10 = v_1 + v_{48} - v_{19} - v_{22} \quad (374)$$

## 9.3 Species S12

**Name** @cyto::p21(b)

**Notes** p21 - Cyclin-dependent kinase inhibitor 1

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in 15 reactions (as a reactant in [reaction\\_22](#), [reaction\\_23](#), [reaction\\_25](#), [reaction\\_26](#), [reaction\\_28](#) and as a product in [reaction\\_5](#), [reaction\\_43](#), [reaction\\_45](#), [reaction\\_48](#), [reaction\\_50](#) and as a modifier in [reaction\\_22](#), [reaction\\_23](#), [reaction\\_25](#), [reaction\\_26](#), [reaction\\_28](#)).

$$\frac{d}{dt}S12 = v_5 + v_{43} + v_{45} + v_{48} + v_{50} - v_{22} - v_{23} - v_{25} - v_{26} - v_{28} \quad (375)$$

## 9.4 Species S19

**Name** @cyto::C4D1(b!1).p21(b!1)

**Notes** Cyclin D1:CDK4:p21 complex

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in nine reactions (as a reactant in [reaction\\_43](#), [reaction\\_48](#), [reaction\\_51](#), [reaction\\_53](#) and as a product in [reaction\\_22](#) and as a modifier in [reaction\\_43](#), [reaction\\_48](#), [reaction\\_51](#), [reaction\\_53](#)).

$$\frac{d}{dt}S19 = v_{22} - v_{43} - v_{48} - v_{51} - v_{53} \quad (376)$$



## 9.5 Species S20

**Name** @cyto::C2E(T160Û,b!1).p21(b!1)

**Notes** Cyclin E:CDK2:p21 complex - T160 phosphorylation

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in nine reactions (as a reactant in [reaction\\_45](#), [reaction\\_50](#), [reaction\\_52](#), [reaction\\_54](#) and as a product in [reaction\\_23](#) and as a modifier in [reaction\\_45](#), [reaction\\_50](#), [reaction\\_52](#), [reaction\\_54](#)).

$$\frac{d}{dt}S20 = v_{23} - v_{45} - v_{50} - v_{52} - v_{54} \quad (377)$$

## 9.6 Species hgf

**Name** HGF

**Notes** Hepatocyte Growth Factor

**Initial concentration** 0 nmol · l<sup>-1</sup>

**Involved in rule** [hgf](#)

One rule determines the species' quantity.

## 9.7 Species inhp53

**Name** inhp53

**Notes** p53 tumor suppressor

**Initial concentration** 0 nmol · l<sup>-1</sup>

**Involved in rule** [inhp53](#)

One rule determines the species' quantity.

## 9.8 Species inherk

**Name** inhERK

**Notes** Extracellular-signal-regulated Kinases

**Initial concentration** 0 nmol · l<sup>-1</sup>

**Involved in rule** [inherk](#)

One rule determines the species' quantity.

### 9.9 Species [inhakt](#)

**Name** inhAKT

**Notes** Protein kinase B

**Initial concentration**  $0 \text{ nmol} \cdot \text{l}^{-1}$

**Involved in rule** [inhakt](#)

One rule determines the species' quantity.

### 9.10 Species [inhc4d1](#)

**Name** inhC4d1

**Notes** Cyclin D1:CDK4 complex

**Initial concentration**  $0 \text{ nmol} \cdot \text{l}^{-1}$

**Involved in rule** [inhc4d1](#)

One rule determines the species' quantity.

### 9.11 Species [ObsTotCycECDK2\\_obs](#)

**Name** TotCycECDK2

**Notes** Cyclin E:CDK2 complex

**Initial concentration**  $0.099620260255418 \text{ nmol} \cdot \text{l}^{-1}$

**Involved in rule** [ObsTotCycECDK2\\_obs](#)

One rule determines the species' quantity.

### 9.12 Species [ObsTotCDK2T160\\_obs](#)

**Name** TotCDK2T160

**Notes** CDK2 T160-phosphorylated (Thr160 site)

**Initial concentration**  $0 \text{ nmol} \cdot \text{l}^{-1}$

**Involved in rule** [ObsTotCDK2T160\\_obs](#)

One rule determines the species' quantity.

### 9.13 Species [ObsTotCycDCDK4\\_obs](#)

**Name** TotCycDCDK4

**Notes** Cyclin D1:CDK4 complex

**Initial concentration** 0 nmol · l<sup>-1</sup>

**Involved in rule** [ObsTotCycDCDK4\\_obs](#)

One rule determines the species' quantity.

### 9.14 Species [ObsTotP21\\_obs](#)

**Name** TotP21

**Notes** p21 - Cyclin-dependent kinase inhibitor 1

**Initial concentration** 0.020805213622291 nmol · l<sup>-1</sup>

**Involved in rule** [ObsTotP21\\_obs](#)

One rule determines the species' quantity.

### 9.15 Species [ObsCDK2P21\\_obs](#)

**Name** CDK2P21

**Notes** p21:CDK2 complex

**Initial concentration** 0.0409109861876772 nmol · l<sup>-1</sup>

**Involved in rule** [ObsCDK2P21\\_obs](#)

One rule determines the species' quantity.

### 9.16 Species [ObsTotE2F\\_obs](#)

**Name** TotE2F

**Notes** E2F-1 - E2F transcription factor

**Initial concentration** 0.0337275199303406 nmol · l<sup>-1</sup>

**Involved in rule** [ObsTotE2F\\_obs](#)

One rule determines the species' quantity.

### 9.17 Species [ObsTotRb\\_obs](#)

**Name** TotRb

**Notes** Retinoblastoma Protein

**Initial concentration**  $0.129299840363777 \text{ nmol} \cdot \text{l}^{-1}$

**Involved in rule** [ObsTotRb\\_obs](#)

One rule determines the species' quantity.

### 9.18 Species [ObsPhosRbS788\\_obs](#)

**Name** PhosRbS788

**Notes** Rb S788-phosphorylated (serine 788 site)

**Initial concentration**  $0 \text{ nmol} \cdot \text{l}^{-1}$

**Involved in rule** [ObsPhosRbS788\\_obs](#)

One rule determines the species' quantity.

### 9.19 Species [ObsPhosRbS800\\_obs](#)

**Name** PhosRbS800

**Notes** Rb S800/S804-phosphorylated (serine 800 and 804 sites)

**Initial concentration**  $0 \text{ nmol} \cdot \text{l}^{-1}$

**Involved in rule** [ObsPhosRbS800\\_obs](#)

One rule determines the species' quantity.

### 9.20 Species [ObsDNAContent\\_obs](#)

**Name** DNAContent

**Notes** DNA content

**Initial concentration**  $1 \text{ nmol} \cdot \text{l}^{-1}$

**Involved in rule** [ObsDNAContent\\_obs](#)

One rule determines the species' quantity.

### 9.21 Species S23

**Name** @nuc::C2E(T160P̃,b!1).p21(b!1)

**Notes** Cyclin E:CDK2:p21 complex - T160 phosphorylation

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in seven reactions (as a reactant in [reaction\\_60](#), [reaction\\_62](#), [reaction\\_63](#) and as a product in [reaction\\_49](#) and as a modifier in [reaction\\_60](#), [reaction\\_62](#), [reaction\\_63](#)).

$$\frac{d}{dt}S23 = v_{49} - v_{60} - v_{62} - v_{63} \quad (378)$$

### 9.22 Species S18

**Name** @nuc::C2E(T160P̃,b)

**Notes** Cyclin E:CDK2 complex - T160 phosphorylation

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in 20 reactions (as a reactant in [reaction\\_44](#), [reaction\\_46](#), [reaction\\_47](#), [reaction\\_49](#) and as a product in [reaction\\_21](#), [reaction\\_62](#), [reaction\\_63](#) and as a modifier in [reaction\\_7](#), [reaction\\_18](#), [reaction\\_38](#), [reaction\\_39](#), [reaction\\_40](#), [reaction\\_41](#), [reaction\\_44](#), [reaction\\_46](#), [reaction\\_47](#), [reaction\\_49](#), [reaction\\_57](#), [reaction\\_63](#), [reaction\\_64](#)).

$$\frac{d}{dt}S18 = v_{21} + v_{62} + v_{63} - v_{44} - v_{46} - v_{47} - v_{49} \quad (379)$$

### 9.23 Species S3

**Name** @nuc::C2E(T160Û,b!1).p21(b!1)

**Notes** Cyclin E:CDK2:p21 complex - T160 phosphorylation

**Initial concentration** 6.2223 nmol · l<sup>-1</sup>

This species takes part in eight reactions (as a reactant in [reaction\\_3](#), [reaction\\_6](#), [reaction\\_7](#) and as a product in [reaction\\_24](#), [reaction\\_52](#) and as a modifier in [reaction\\_3](#), [reaction\\_6](#), [reaction\\_7](#)).

$$\frac{d}{dt}S3 = v_{24} + v_{52} - v_3 - v_6 - v_7 \quad (380)$$

## 9.24 Species S13

**Name** @nuc::C2E(T160Û,b)

**Notes** Cyclin E:CDK2 complex - T160 phosphorylation

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in nine reactions (as a reactant in [reaction\\_20](#), [reaction\\_21](#), [reaction\\_24](#) and as a product in [reaction\\_6](#), [reaction\\_7](#), [reaction\\_47](#) and as a modifier in [reaction\\_20](#), [reaction\\_21](#), [reaction\\_24](#)).

$$\frac{d}{dt}S13 = v_6 + v_7 + v_{47} - v_{20} - v_{21} - v_{24} \quad (381)$$

## 9.25 Species S24

**Name** @nuc::C4D1(b!1),p21(b!1)

**Notes** Cyclin D1:CDK4:p21 complex

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in ten reactions (as a reactant in [reaction\\_59](#), [reaction\\_61](#), [reaction\\_64](#) and as a product in [reaction\\_51](#), [reaction\\_67](#) and as a modifier in [reaction\\_16](#), [reaction\\_17](#), [reaction\\_59](#), [reaction\\_61](#), [reaction\\_64](#)).

$$\frac{d}{dt}S24 = v_{51} + v_{67} - v_{59} - v_{61} - v_{64} \quad (382)$$

## 9.26 Species S26

**Name** @nuc::C4D1(b)

**Notes** Cyclin D1:CDK4 complex

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in six reactions (as a reactant in [reaction\\_66](#), [reaction\\_67](#) and as a product in [reaction\\_61](#), [reaction\\_64](#) and as a modifier in [reaction\\_66](#), [reaction\\_67](#)).

$$\frac{d}{dt}S26 = v_{61} + v_{64} - v_{66} - v_{67} \quad (383)$$

### 9.27 Species S5

**Name** @nuc::dnapre()

**Notes** Genes contributing to the formation of prereplication complexes.

**Initial concentration** 1 nmol · l<sup>-1</sup>

This species takes part in two reactions (as a reactant in [reaction\\_18](#) and as a modifier in [reaction\\_18](#)).

$$\frac{d}{dt}S5 = -v_{18} \quad (384)$$

### 9.28 Species S17

**Name** @nuc::dnapre1()

**Notes** Genes contributing to the formation of prereplication complexes.

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in three reactions (as a reactant in [reaction\\_42](#) and as a product in [reaction\\_18](#) and as a modifier in [reaction\\_42](#)).

$$\frac{d}{dt}S17 = v_{18} - v_{42} \quad (385)$$

### 9.29 Species S22

**Name** @nuc::dnapre2()

**Notes** Genes contributing to the formation of prereplication complexes.

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in three reactions (as a reactant in [reaction\\_58](#) and as a product in [reaction\\_42](#) and as a modifier in [reaction\\_58](#)).

$$\frac{d}{dt}S22 = v_{42} - v_{58} \quad (386)$$

### 9.30 Species S25

**Name** @nuc::dnapre3()

**Notes** Genes contributing to the formation of prereplication complexes.

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in three reactions (as a reactant in [reaction\\_65](#) and as a product in [reaction\\_58](#) and as a modifier in [reaction\\_65](#)).

$$\frac{d}{dt}S25 = v_{58} - v_{65} \quad (387)$$

### 9.31 Species S27

**Name** @nuc::dnapre4()

**Notes** Genes contributing to the formation of prereplication complexes.

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in three reactions (as a reactant in [reaction\\_68](#) and as a product in [reaction\\_65](#) and as a modifier in [reaction\\_68](#)).

$$\frac{d}{dt}S27 = v_{65} - v_{68} \quad (388)$$

### 9.32 Species S16

**Name** @nuc::e2f(b!1).rb(S788P̃,S800Ũ,b!1)

**Notes** E2F-1: Retinoblastoma Protein complex - S788 and S800 phosphorylation

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in 15 reactions (as a reactant in [reaction\\_30](#), [reaction\\_32](#), [reaction\\_34](#), [reaction\\_37](#), [reaction\\_39](#), [reaction\\_41](#) and as a product in [reaction\\_17](#), [reaction\\_36](#) and as a modifier in [reaction\\_2](#), [reaction\\_30](#), [reaction\\_32](#), [reaction\\_34](#), [reaction\\_37](#), [reaction\\_39](#), [reaction\\_41](#)).

$$\frac{d}{dt}S16 = v_{17} + v_{36} - v_{30} - v_{32} - v_{34} - v_{37} - v_{39} - v_{41} \quad (389)$$

### 9.33 Species S2

**Name** @nuc::e2f(b!1).rb(S788Ũ,S800Ũ,b!1)

**Notes** E2F-1: Retinoblastoma Protein complex - S788 and S800 phosphorylation

**Initial concentration** 0.0601 nmol · l<sup>-1</sup>

This species takes part in twelve reactions (as a reactant in [reaction\\_10](#), [reaction\\_12](#), [reaction\\_14](#), [reaction\\_15](#), [reaction\\_17](#) and as a product in [reaction\\_35](#), [reaction\\_41](#) and as a modifier in [reaction\\_10](#), [reaction\\_12](#), [reaction\\_14](#), [reaction\\_15](#), [reaction\\_17](#)).

$$\frac{d}{dt}S2 = v_{35} + v_{41} - v_{10} - v_{12} - v_{14} - v_{15} - v_{17} \quad (390)$$



### 9.34 Species S14

**Name** @nuc::e2f(b)

**Notes** E2F-1 - E2F transcription factor

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in 22 reactions (as a reactant in [reaction\\_33](#), [reaction\\_35](#), [reaction\\_36](#) and as a product in [reaction\\_10](#), [reaction\\_12](#), [reaction\\_13](#), [reaction\\_15](#), [reaction\\_30](#), [reaction\\_32](#), [reaction\\_37](#), [reaction\\_39](#) and as a modifier in [reaction\\_2](#), [reaction\\_5](#), [reaction\\_7](#), [reaction\\_8](#), [reaction\\_13](#), [reaction\\_18](#), [reaction\\_33](#), [reaction\\_35](#), [reaction\\_36](#), [reaction\\_63](#), [reaction\\_64](#)).

$$\frac{d}{dt}S14 = v_{10} + v_{12} + v_{13} + v_{15} + v_{30} + v_{32} + v_{37} + v_{39} - v_{33} - v_{35} - v_{36} \quad (391)$$

### 9.35 Species S11

**Name** @nuc::p21(b)

**Notes** p21 - Cyclin-dependent kinase inhibitor 1

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in 19 reactions (as a reactant in [reaction\\_24](#), [reaction\\_27](#), [reaction\\_49](#), [reaction\\_67](#) and as a product in [reaction\\_3](#), [reaction\\_6](#), [reaction\\_25](#), [reaction\\_60](#), [reaction\\_61](#), [reaction\\_62](#) and as a modifier in [reaction\\_11](#), [reaction\\_12](#), [reaction\\_24](#), [reaction\\_27](#), [reaction\\_31](#), [reaction\\_32](#), [reaction\\_49](#), [reaction\\_56](#), [reaction\\_67](#)).

$$\frac{d}{dt}S11 = v_3 + v_6 + v_{25} + v_{60} + v_{61} + v_{62} - v_{24} - v_{27} - v_{49} - v_{67} \quad (392)$$

### 9.36 Species S21

**Name** @nuc::rb(S788P̃,S800P̃,b)

**Notes** Retinoblastoma Protein - S788 and S800 phosphorylation

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in eight reactions (as a reactant in [reaction\\_55](#), [reaction\\_56](#), [reaction\\_57](#) and as a product in [reaction\\_38](#), [reaction\\_39](#) and as a modifier in [reaction\\_55](#), [reaction\\_56](#), [reaction\\_57](#)).

$$\frac{d}{dt}S21 = v_{38} + v_{39} - v_{55} - v_{56} - v_{57} \quad (393)$$

### 9.37 Species S15

**Name** @nuc::rb(S788P̃,S800Ũ,b)

**Notes** Retinoblastoma Protein - S788 and S800 phosphorylation

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in 14 reactions (as a reactant in [reaction\\_29](#), [reaction\\_31](#), [reaction\\_36](#), [reaction\\_38](#), [reaction\\_40](#) and as a product in [reaction\\_16](#), [reaction\\_34](#), [reaction\\_37](#), [reaction\\_57](#) and as a modifier in [reaction\\_29](#), [reaction\\_31](#), [reaction\\_36](#), [reaction\\_38](#), [reaction\\_40](#)).

$$\frac{d}{dt}S15 = v_{16} + v_{34} + v_{37} + v_{57} - v_{29} - v_{31} - v_{36} - v_{38} - v_{40} \quad (394)$$

### 9.38 Species S1

**Name** @nuc::rb(S788Ũ,S800Ũ,b)

**Notes** Retinoblastoma Protein - S788 and S800 phosphorylation

**Initial concentration** 25.5914 nmol · l<sup>-1</sup>

This species takes part in twelve reactions (as a reactant in [reaction\\_9](#), [reaction\\_11](#), [reaction\\_16](#), [reaction\\_35](#) and as a product in [reaction\\_8](#), [reaction\\_14](#), [reaction\\_15](#), [reaction\\_40](#) and as a modifier in [reaction\\_9](#), [reaction\\_11](#), [reaction\\_16](#), [reaction\\_35](#)).

$$\frac{d}{dt}S1 = v_8 + v_{14} + v_{15} + v_{40} - v_9 - v_{11} - v_{16} - v_{35} \quad (395)$$

### 9.39 Species S28

**Name** @nuc::dnapre5()

**Notes** Genes contributing to the formation of prereplication complexes.

**Initial concentration** 0 nmol · l<sup>-1</sup>

This species takes part in three reactions (as a reactant in [reaction\\_69](#) and as a product in [reaction\\_68](#) and as a modifier in [reaction\\_69](#)).

$$\frac{d}{dt}S28 = v_{68} - v_{69} \quad (396)$$

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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