# **SBML Model Report**

# Model name: "Leloup1999\_CircClock"



August 10, 2016

# 1 General Overview

This is a document in SBML Level 2 Version 1 format. This model was created by the following two authors: Nicolas Le Novre<sup>1</sup> and Bruce Shapiro<sup>2</sup> at June 29<sup>th</sup> 2005 at 10:27 a.m. and last time modified at February 25<sup>th</sup> 2015 at 1:16 p.m. Table 1 provides an overview of the quantities of all components of this model.

Table 1: Number of components in this model, which are described in the following sections.

Element	Quantity	Element	Quantity
compartment types	0	compartments	2
species types	0	species	10
events	0	constraints	0
reactions	24	function definitions	0
global parameters	4	unit definitions	2
rules	2	initial assignments	0

#### **Model Notes**

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To cite BioModels Database, please use Le Novre N., Bornstein B., Broicher A., Courtot M., Donizelli M., Dharuri H., Li L., Sauro H., Schilstra M., Shapiro B., Snoep J.L., Hucka M. (2006)

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BioModels Database: A Free, Centralized Database of Curated, Published, Quantitative Kinetic Models of Biochemical and Cellular Systems Nucleic Acids Res., 34: D689-D691.

# **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 substance

Name nanomole (default)

**Definition** nmol

#### 2.2 Unit time

Name hour (default)

**Definition** 3600 s

#### 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** m<sup>2</sup>

# 2.5 Unit length

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

**Definition** m

# 3 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		
compartment_0000002	nucleus		3	1	litre		Cell

# 3.1 Compartment Cell

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

Name cytoplasm

# **3.2 Compartment** compartment\_0000002

This is a three dimensional compartment with a constant size of one litre, which is surrounded by Cell (cytoplasm).

Name nucleus

# Produced by SBML2ATEX

# 4 Species

This model contains ten species. Section 8 provides further details and the derived rates of change of each species.

Table 3: Properties of each species.

Id	Name	Compartment	Derived Unit	Constant	Boundary Condi- tion
PO	PER Protein (unphosphorylated)	Cell	$nmol \cdot l^{-1}$		
TO	TIM Protein (unphosphorylated)	Cell	$nmol \cdot l^{-1}$		$\Box$
P1	PER Protein (mono-phosphorylated)	Cell	$nmol \cdot l^{-1}$		$\Box$
T1	TIM Protein (mono-phosphorylated)	Cell	$nmol \cdot l^{-1}$		$\Box$
P2	PER Protein (bi-phosphorylated)	Cell	$nmol \cdot l^{-1}$		$\Box$
T2	TIM Protein (bi-phosphorylated)	Cell	$nmol \cdot l^{-1}$		$\Box$
CC	Cytosolic PER-TIM Complex	Cell	$nmol \cdot l^{-1}$		$\Box$
Cn	Nuclear PER-TIM Complex	$compartment_0000002$	$nmol \cdot l^{-1}$		$\Box$
Mp	PER mRNA	Cell	$nmol \cdot l^{-1}$		$\Box$
Mt	TIM mRNA	Cell	$nmol \cdot l^{-1}$		

# **5 Parameters**

This model contains four global parameters.

Table 4: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
Pt	Total Per	0.0	
Tt	Total Tim	0.0	
$V_mT$		0.7	$\square$
$V_{-}dT$		2.0	$\checkmark$

# 6 Rules

This is an overview of two rules.

# 6.1 Rule Pt

Rule Pt is an assignment rule for parameter Pt:

$$Pt = [CC] + [Cn] + [P0] + [P1] + [P2]$$
 (1)

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

# 6.2 Rule Tt

Rule Tt is an assignment rule for parameter Tt:

$$Tt = [CC] + [Cn] + [T0] + [T1] + [T2]$$
 (2)

**Derived unit**  $nmol \cdot l^{-1}$ 

# 7 Reactions

This model contains 24 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	PO_to_P1	First Phosphorylation of PER	P0 → P1	
2	TO_to_T1	First Phosphorylation of TIM	$T0 \longrightarrow T1$	
3	P1_to_P0	Dephosphorylation of PER (1st P)	$P1 \longrightarrow P0$	
4	$T1_to_T0$	Dephosphorylation of TIM (1st P)	$T1 \longrightarrow T0$	
5	P1_to_P2	Second Phosphorylation of PER	$P1 \longrightarrow P2$	
6	$T1_to_T2$	Second Phosphorylation of TIM	$T1 \longrightarrow T2$	
7	P2_to_P1	Dephosphorylation of PER (2nd P)	$P2 \longrightarrow P1$	
8	T2_to_T1	Dephosphorylation of TIM (2nd P)	$T2 \longrightarrow T1$	
9	$PO_{-}$ degradation	PER degradation	$P0 \longrightarrow \emptyset$	
10	${\tt T0\_degradation}$	TIM degradation	$T0 \longrightarrow \emptyset$	
11	$P1\_degradation$	PER-1 degradation	$P1 \longrightarrow \emptyset$	
12	$T1\_degradation$	TIM-1 degradation	$T1 \longrightarrow \emptyset$	
13	$P2\_degradation$	PER-2 degradation	$P2 \longrightarrow \emptyset$	
14	$T2\_degradation$	TIM-2 degradation	$T2 \longrightarrow \emptyset$	
15	PT_complex- _formation	PER-TIM complex formation	$P2 + T2 \Longrightarrow CC$	
16	PT_complex- _nucleation	PER-TIM complex nucleation	CC <del>←</del> Cn	
17	PT_complexdegradation	PER-TIM complex degradation (cytosol)	$CC \longrightarrow \emptyset$	
18	PTnucl_complex- _degradation	PER-TIM complex degradation (nuclear)	$Cn \longrightarrow \emptyset$	

N⁰	Id	Name	Reaction Equation	SBO
19	Mp_production	PER mRNA production	$\emptyset \xrightarrow{Cn} Mp$	
20	$\mathtt{Mt\_production}$	TIM mRNA production	$\emptyset \xrightarrow{Cn} Mt$	
21	$PO_{-}$ production	PER production	$\emptyset \xrightarrow{Mp} P0$	
22	$T0_production$	TIM production	$\emptyset \xrightarrow{\mathbf{Mt}} \mathbf{T0}$	
23	$\mathtt{Mp\_degradation}$	PER mRNA degradation	$Mp \longrightarrow \emptyset$	
24	${\tt Mt\_degradation}$	TIM mRNA degradation	$Mt \longrightarrow \emptyset$	

# **7.1 Reaction PO\_to\_P1**

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

Name First Phosphorylation of PER

**Notes** This phosphorylation is triggered by the protein product of the gene <u>double-time</u> (DBT, <u>DCO\_DROME</u>). Not explicitly represented in the model.

# **Reaction equation**

$$P0 \longrightarrow P1$$
 (3)

#### Reactant

Table 6: Properties of each reactant.

Id	Name	SBO
PO	PER Protein (unphosphorylated)	

# **Product**

Table 7: Properties of each product.

Id	Name	SBO
P1	PER Protein (mono-phosphorylated)	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_1 = \frac{\text{vol}\left(\text{Cell}\right) \cdot \text{V}_{-}1\text{P} \cdot [\text{P0}]}{\text{K1}_{-}\text{P} + [\text{P0}]} \tag{4}$$

Table 8: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
K1_P		2.0	
$V_{-}1P$		8.0	$\square$

# 7.2 Reaction TO\_to\_T1

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

Name First Phosphorylation of TIM

# **Reaction equation**

$$T0 \longrightarrow T1$$
 (5)

# Reactant

Table 9: Properties of each reactant.

Id	Name	SBO
ТО	TIM Protein (unphosphorylated)	

# **Product**

Table 10: Properties of each product.

Id	Name	SBO
T1	TIM Protein (mono-phosphorylated)	

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_2 = \frac{\text{vol}(\text{Cell}) \cdot \text{V}_{-}1\text{T} \cdot [\text{T0}]}{\text{K}_{-}1\text{T} + [\text{T0}]}$$
(6)

Table 11: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
K_1T		2.0	
$V_{-}1T$		8.0	$\checkmark$

# **7.3 Reaction P1\_to\_P0**

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

Name Dephosphorylation of PER (1st P)

# **Reaction equation**

$$P1 \longrightarrow P0$$
 (7)

# Reactant

Table 12: Properties of each reactant.

	rue re reperires or euch reuctum.	
Id	Name	SBO
P1	PER Protein (mono-phosphorylated)	

# **Product**

Table 13: Properties of each product.

	Tueste Text Treperines of each product	
Id	Name	SBO
P0	PER Protein (unphosphorylated)	

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_3 = \frac{\text{vol}(\text{Cell}) \cdot \text{V}_2\text{P} \cdot [\text{P1}]}{\text{K}_2\text{P} + [\text{P1}]}$$
(8)

Table 14: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
K_2P		2.0	$\overline{Z}$
$V_2P$		1.0	

# 7.4 Reaction T1\_to\_T0

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

Name Dephosphorylation of TIM (1st P)

# **Reaction equation**

$$T1 \longrightarrow T0$$
 (9)

#### Reactant

Table 15: Properties of each reactant.

Id	Name	SBO
T1	TIM Protein (mono-phosphorylated)	

# **Product**

Table 16: Properties of each product.

Id	Name	SBO
ТО	TIM Protein (unphosphorylated)	

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_4 = \frac{\text{vol}(\text{Cell}) \cdot \text{V}.2\text{T} \cdot [\text{T1}]}{\text{K}.2\text{T} + [\text{T1}]}$$
 (10)

Table 17: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
K_2T		2.0	$ \mathbf{Z} $
$V_2T$		1.0	$\overline{\mathbb{Z}}$

# 7.5 Reaction P1\_to\_P2

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

Name Second Phosphorylation of PER

**Notes** This phosphorylation is triggered by the protein product of the gene <u>double-time</u> (DBT, <u>DCO\_DROME</u>). Not explicitly represented in the model.

# **Reaction equation**

$$P1 \longrightarrow P2$$
 (11)

# Reactant

Table 18: Properties of each reactant.

Table 10. 110perties of each reactant.			
Id	Name	SBO	
P1	PER Protein (mono-phosphorylated)		

Table 19: Properties of each product.

Id	Name	SBO
P2	PER Protein (bi-phosphorylated)	

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_5 = \frac{\text{vol}(\text{Cell}) \cdot \text{V}_{-3}\text{P} \cdot [\text{P1}]}{\text{K}_{-3}\text{P} + [\text{P1}]}$$
(12)

Table 20: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
K_3P		2.0	
$V_3P$		8.0	

# 7.6 Reaction T1\_to\_T2

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

Name Second Phosphorylation of TIM

# **Reaction equation**

$$T1 \longrightarrow T2$$
 (13)

# Reactant

Table 21: Properties of each reactant.

Id	Name	SBO
T1	TIM Protein (mono-phosphorylated)	

Table 22: Properties of each product.

Id	Name	SBO
T2	TIM Protein (bi-phosphorylated)	

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_6 = \frac{\text{vol}(\text{Cell}) \cdot \text{V}_{-3}\text{T} \cdot [\text{T1}]}{\text{K}_{-3}\text{T} + [\text{T1}]}$$
(14)

Table 23: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
K_3T		2.0	
$V_{-}3T$		8.0	

# 7.7 Reaction P2\_to\_P1

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

Name Dephosphorylation of PER (2nd P)

# **Reaction equation**

$$P2 \longrightarrow P1$$
 (15)

# Reactant

Table 24: Properties of each reactant.

Id	Name	SBO
P2	PER Protein (bi-phosphorylated)	

# **Product**

Table 25: Properties of each product.

	P P	
Id	Name	SBO
P1	PER Protein (mono-phosphorylated)	

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_7 = \frac{\text{vol}\left(\text{Cell}\right) \cdot \text{V}_{\text{-}}4\text{P} \cdot [\text{P2}]}{\text{K}_{\text{-}}4\text{P} + [\text{P2}]} \tag{16}$$

Table 26: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
K_4P		2.0	
$V_{-}4P$		1.0	$\square$

# 7.8 Reaction T2\_to\_T1

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

Name Dephosphorylation of TIM (2nd P)

# **Reaction equation**

$$T2 \longrightarrow T1$$
 (17)

#### Reactant

Table 27: Properties of each reactant.

Id	Name	SBO
T2	TIM Protein (bi-phosphorylated)	

# **Product**

Table 28: Properties of each product.

Id	Name	SBO
T1	TIM Protein (mono-phosphorylated)	

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_8 = \frac{\text{vol}(\text{Cell}) \cdot \text{V}_{-}4\text{T} \cdot [\text{T2}]}{\text{K}_{-}4\text{T} + [\text{T2}]}$$
(18)

Table 29: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
K_4T		2.0	$\overline{Z}$
$V_{-}4T$		1.0	$\checkmark$

# **7.9 Reaction** PO\_degradation

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

Name PER degradation

# **Reaction equation**

$$P0 \longrightarrow \emptyset \tag{19}$$

#### Reactant

Table 30: Properties of each reactant.

Tuble 30. I roperties of each reactant.			
Id	Name	SBO	
P0	PER Protein (unphosphorylated)		

# **Kinetic Law**

Derived unit contains undeclared units

$$v_9 = \text{vol}(\text{Cell}) \cdot \text{k\_d} \cdot [\text{P0}] \tag{20}$$

Table 31: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k_d		0.01	

# **7.10 Reaction** TO\_degradation

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

Name TIM degradation

# **Reaction equation**

$$T0 \longrightarrow \emptyset$$
 (21)

# Reactant

Table 32: Properties of each reactant.

	I	
Id	Name	SBO
TO	TIM Protein (unphosphorylated)	

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{10} = \text{vol}\left(\text{Cell}\right) \cdot \text{k\_d} \cdot [\text{T0}] \tag{22}$$

Table 33: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k_d		0.01	

# 7.11 Reaction P1\_degradation

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

Name PER-1 degradation

# **Reaction equation**

$$P1 \longrightarrow \emptyset \tag{23}$$

# Reactant

Table 34: Properties of each reactant.

Table 34. I Toperties of each reactant.			
Id	Name	SBO	
P1	PER Protein (mono-phosphorylated)		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{11} = \text{vol}\left(\text{Cell}\right) \cdot k_{-}d \cdot [\text{P1}] \tag{24}$$

Table 35: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k_d		0.01	$\overline{Z}$

# **7.12 Reaction** T1\_degradation

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

Name TIM-1 degradation

# **Reaction equation**

$$T1 \longrightarrow \emptyset$$
 (25)

# Reactant

Table 36: Properties of each reactant.

Id	Name	SBO
T1	TIM Protein (mono-phosphorylated)	

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{12} = \text{vol}\left(\text{Cell}\right) \cdot k_{-}d \cdot [\text{T1}] \tag{26}$$

Table 37: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k_d		0.01	$\checkmark$

# 7.13 Reaction P2\_degradation

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

Name PER-2 degradation

# **Reaction equation**

$$P2 \longrightarrow \emptyset \tag{27}$$

#### Reactant

Table 38: Properties of each reactant.

Id	Name	SBO
P2	PER Protein (bi-phosphorylated)	

# **Kinetic Law**

Derived unit contains undeclared units

$$v_{13} = vol\left(Cell\right) \cdot k\_d \cdot [P2] + \frac{vol\left(Cell\right) \cdot V\_dP \cdot [P2]}{K\_dP + [P2]}$$
 (28)

Table 39: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
$k_{-}d$		0.01	$\square$
$V_{-}dP$		2.00	$ \overline{\checkmark} $
$K_{-}dP$		0.20	

# **7.14 Reaction** T2\_degradation

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

Name TIM-2 degradation

# **Reaction equation**

$$T2 \longrightarrow \emptyset$$
 (29)

#### Reactant

Table 40: Properties of each reactant.

Id	Name	SBO
T2	TIM Protein (bi-phosphorylated)	

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{14} = \text{vol}\left(\text{Cell}\right) \cdot \text{k\_d} \cdot [\text{T2}] + \frac{\text{vol}\left(\text{Cell}\right) \cdot \text{V\_dT} \cdot [\text{T2}]}{\text{K\_dT} + [\text{T2}]}$$
(30)

Table 41: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k_d		0.01	
$K_dT$		0.20	$\overline{\checkmark}$

# **7.15 Reaction PT\_complex\_formation**

This is a reversible reaction of two reactants forming one product.

Name PER-TIM complex formation

# **Reaction equation**

$$P2 + T2 \Longrightarrow CC$$
 (31)

# Reactants

Table 42: Properties of each reactant.

Id	Name	SBO
P2	PER Protein (bi-phosphorylated)	
T2	TIM Protein (bi-phosphorylated)	

Table 43: Properties of each product.

	Name	SBO
CC	Cytosolic PER-TIM Complex	

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{15} = \text{vol}(\text{Cell}) \cdot \text{k3} \cdot [\text{P2}] \cdot [\text{T2}] - \text{vol}(\text{Cell}) \cdot \text{k4} \cdot [\text{CC}]$$
(32)

Table 44: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k3		1.2	
k4		0.6	$\checkmark$

# **7.16 Reaction** PT\_complex\_nucleation

This is a reversible reaction of one reactant forming one product.

Name PER-TIM complex nucleation

# **Reaction equation**

$$CC \rightleftharpoons Cn$$
 (33)

# Reactant

Table 45: Properties of each reactant.

Id	Name	SBO
CC	Cytosolic PER-TIM Complex	

# **Product**

Table 46: Properties of each product

Table 40. I toperties of each product.			
Id	Name	SBO	
Cn	Nuclear PER-TIM Complex		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{16} = vol\left(Cell\right) \cdot k1 \cdot [CC] - vol\left(compartment\_0000002\right) \cdot k2 \cdot [Cn] \tag{34}$$

Table 47: Properties of each parameter.

			•	
Id	Name	SBO Va	lue Unit	Constant
k1		0	.6	
k2		0.	.2	

# **7.17 Reaction PT\_complex\_degradation**

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

Name PER-TIM complex degradation (cytosol)

# **Reaction equation**

$$CC \longrightarrow \emptyset$$
 (35)

#### Reactant

Table 48: Properties of each reactant.

	Name	SBO
CC	Cytosolic PER-TIM Complex	

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{17} = \text{vol}(\text{Cell}) \cdot \text{k\_dC} \cdot [\text{CC}]$$
(36)

Table 49: Properties of each parameter.

Id	Name	SBO	Value	Unit	Constant
k_dC			0.01		

# 7.18 Reaction PTnucl\_complex\_degradation

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

Name PER-TIM complex degradation (nuclear)

# **Reaction equation**

$$Cn \longrightarrow \emptyset$$
 (37)

#### Reactant

Table 50: Properties of each reactant.

Id	Name	SBO
Cn	Nuclear PER-TIM Complex	

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{18} = \text{vol} \left( \text{compartment\_0000002} \right) \cdot \text{k\_dN} \cdot [\text{Cn}]$$
 (38)

Table 51: Properties of each parameter.

Id	Name	SBO Value U	nit Constant
k_dN		0.01	

# 7.19 Reaction Mp\_production

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

Name PER mRNA production

# **Reaction equation**

$$\emptyset \xrightarrow{\operatorname{Cn}} \operatorname{Mp} \tag{39}$$

#### **Modifier**

Table 52: Properties of each modifier.

ruote 32. I roperties of euch mounter.				
Id	Name	SBO		
Cn	Nuclear PER-TIM Complex			

# **Product**

Table 53: Properties of each product.

Id	Name	SBO
Мр	PER mRNA	_

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{19} = \frac{\text{vol}\left(\text{Cell}\right) \cdot \text{v\_sP} \cdot \text{K\_IP}^{\text{n}}}{\text{K\_IP}^{\text{n}} + [\text{Cn}]^{\text{n}}}$$
(40)

Table 54: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
v_sP		1.0	$\square$
$K_{-}IP$		1.0	
n		4.0	$\square$

# **7.20 Reaction Mt\_production**

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

Name TIM mRNA production

# **Reaction equation**

$$\emptyset \xrightarrow{Cn} Mt$$
 (41)

#### **Modifier**

Table 55: Properties of each modifier.

Table 33. I roperties of each infounct.				
Id	Name	SBO		
Cn	Nuclear PER-TIM Complex			

Table 56: Properties of each product.

Id	Name	SBO
Mt	TIM mRNA	

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{20} = \frac{\text{vol}\left(\text{Cell}\right) \cdot \text{V}_{-\text{S}}\text{T} \cdot \text{K}_{-\text{I}}\text{T}^{\text{n}}}{\text{K}_{-\text{I}}\text{T}^{\text{n}} + [\text{Cn}]^{\text{n}}}$$
(42)

Table 57: Properties of each parameter.

		•	•		
Id	Name	SBO	Value	Unit	Constant
V_sT			1.0		
$K_{-}IT$			1.0		$\mathbf{Z}$
n			4.0		$\square$

# 7.21 Reaction PO\_production

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

Name PER production

# **Reaction equation**

$$\emptyset \xrightarrow{Mp} P0$$
 (43)

# **Modifier**

Table 58: Properties of each modifier.

Id	Name	SBO
Мр	PER mRNA	

Table 59: Properties of each product.

	T T T T T T T T T T T T T T T T T T T	
Id	Name	SBO
P0	PER Protein (unphosphorylated)	

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{21} = \text{vol}\left(\text{Cell}\right) \cdot k_{-s}P \cdot [Mp] \tag{44}$$

Table 60: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k_sP		0.9	Ø

# **7.22 Reaction** TO\_production

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

Name TIM production

# **Reaction equation**

$$\emptyset \xrightarrow{Mt} T0$$
 (45)

# **Modifier**

Table 61: Properties of each modifier.

Id	Name	SBO
Mt	TIM mRNA	

Table 62: Properties of each product.

Id	Name	SBO
ТО	TIM Protein (unphosphorylated)	

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{22} = \text{vol}\left(\text{Cell}\right) \cdot \text{k\_sT} \cdot [\text{Mt}] \tag{46}$$

Table 63: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k_sT		0.9	

# 7.23 Reaction Mp\_degradation

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

Name PER mRNA degradation

# **Reaction equation**

$$Mp \longrightarrow \emptyset \tag{47}$$

#### Reactant

Table 64: Properties of each reactant.

Id	Name	SBO
Мр	PER mRNA	

# **Kinetic Law**

Derived unit contains undeclared units

$$v_{23} = \text{vol}\left(\text{Cell}\right) \cdot k\_d \cdot [\text{Mp}] + \frac{\text{vol}\left(\text{Cell}\right) \cdot V\_mP \cdot [\text{Mp}]}{K\_mP + [\text{Mp}]}$$
(48)

Table 65: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k_d		0.01	$ \mathcal{A} $
$V_mP$		0.70	
K_mP		0.20	$\square$

# 7.24 Reaction Mt\_degradation

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

Name TIM mRNA degradation

# **Reaction equation**

$$Mt \longrightarrow \emptyset$$
 (49)

#### Reactant

Table 66: Properties of each reactant.

Id	Name	SBO
Mt	TIM mRNA	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{24} = \text{vol}\left(\text{Cell}\right) \cdot k\_d \cdot [\text{Mt}] + \frac{\text{vol}\left(\text{Cell}\right) \cdot V\_mT \cdot [\text{Mt}]}{K\_mT + [\text{Mt}]}$$
(50)

Table 67: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k_d		0.01	$\overline{Z}$
$K_mT$		0.20	

# 8 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.

# 8.1 Species P0

Name PER Protein (unphosphorylated)

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

This species takes part in four reactions (as a reactant in PO\_to\_P1, PO\_degradation and as a product in P1\_to\_P0, PO\_production).

$$\frac{\mathrm{d}}{\mathrm{d}t}P0 = |v_3| + |v_{21}| - |v_1| - |v_9| \tag{51}$$

# 8.2 Species TO

Name TIM Protein (unphosphorylated)

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

This species takes part in four reactions (as a reactant in T0\_to\_T1, T0\_degradation and as a product in T1\_to\_T0, T0\_production).

$$\frac{\mathrm{d}}{\mathrm{d}t}T0 = |v_4| + |v_{22}| - |v_2| - |v_{10}| \tag{52}$$

#### 8.3 Species P1

Name PER Protein (mono-phosphorylated)

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

This species takes part in five reactions (as a reactant in P1\_to\_P0, P1\_to\_P2, P1\_degradation and as a product in P0\_to\_P1, P2\_to\_P1).

$$\frac{\mathrm{d}}{\mathrm{d}t} P1 = |v_1| + |v_7| - |v_3| - |v_5| - |v_{11}| \tag{53}$$

# 8.4 Species T1

Name TIM Protein (mono-phosphorylated)

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

This species takes part in five reactions (as a reactant in T1\_to\_T0, T1\_to\_T2, T1\_degradation and as a product in T0\_to\_T1, T2\_to\_T1).

$$\frac{\mathrm{d}}{\mathrm{d}t} T1 = v_2 + |v_8| - |v_4| - |v_6| - |v_{12}| \tag{54}$$

# 8.5 Species P2

Name PER Protein (bi-phosphorylated)

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

This species takes part in four reactions (as a reactant in P2\_to\_P1, P2\_degradation, PT\_complex\_formation and as a product in P1\_to\_P2).

$$\frac{\mathrm{d}}{\mathrm{d}t}P2 = |v_5| - |v_7| - |v_{13}| - |v_{15}| \tag{55}$$

# 8.6 Species T2

Name TIM Protein (bi-phosphorylated)

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

This species takes part in four reactions (as a reactant in T2\_to\_T1, T2\_degradation, PT\_complex\_formation and as a product in T1\_to\_T2).

$$\frac{\mathrm{d}}{\mathrm{d}t} T2 = v_6 - |v_8| - |v_{14}| - |v_{15}| \tag{56}$$

# 8.7 Species CC

Name Cytosolic PER-TIM Complex

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

This species takes part in three reactions (as a reactant in PT\_complex\_nucleation, PT\_complex\_degradation and as a product in PT\_complex\_formation).

$$\frac{d}{dt}CC = |v_{15}| - |v_{16}| - |v_{17}| \tag{57}$$

# 8.8 Species Cn

Name Nuclear PER-TIM Complex

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

This species takes part in four reactions (as a reactant in PTnucl\_complex\_degradation and as a product in PT\_complex\_nucleation and as a modifier in Mp\_production, Mt\_production).

$$\frac{d}{dt}Cn = v_{16} - v_{18} \tag{58}$$

# 8.9 Species Mp

Name PER mRNA

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

This species takes part in three reactions (as a reactant in Mp\_degradation and as a product in Mp\_production and as a modifier in P0\_production).

$$\frac{d}{dt}Mp = |v_{19}| - |v_{23}| \tag{59}$$

# 8.10 Species Mt

Name TIM mRNA

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

This species takes part in three reactions (as a reactant in Mt\_degradation and as a product in Mt\_production and as a modifier in T0\_production).

$$\frac{d}{dt}Mt = v_{20} - v_{24} \tag{60}$$

 $\mathfrak{BML2}^{d}$  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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