

## SBML Model Report

### Model name: “Bertram2007\_IsletCell\_Oscillations”



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: Ishan Ajmera<sup>1</sup> and Catherine Lloyd<sup>2</sup> at September 29<sup>th</sup> 2011 at 10:13 p. m. and last time modified at April eighth 2016 at 5:08 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	1
species types	0	species	11
events	0	constraints	0
reactions	0	function definitions	0
global parameters	151	unit definitions	0
rules	94	initial assignments	0

## Model Notes

This is the model described in the article:

**Interaction of glycolysis and mitochondrial respiration in metabolic oscillations of pancreatic islets.**

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Bertram R, Satin LS, Pedersen MG, Luciani DS, Sherman A. *Biophys J.* 2007 Mar 1;92(5):1544-55. Pubmed ID: [17172305](#), doi: [10.1529/biophysj.106.097154](#).

**Abstract:**

Insulin secretion from pancreatic beta-cells is oscillatory, with a typical period of 2-7 min, reflecting oscillations in membrane potential and the cytosolic Ca(2+) concentration. Our central hypothesis is that the slow 2-7 min oscillations are due to glycolytic oscillations, whereas faster oscillations that are superimposed are due to Ca(2+) feedback onto metabolism or ion channels. We extend a previous mathematical model based on this hypothesis to include a more detailed description of mitochondrial metabolism. We demonstrate that this model can account for typical oscillatory patterns of membrane potential and Ca(2+) concentration in islets. It also accounts for temporal data on oxygen consumption in islets. A recent challenge to the notion that glycolytic oscillations drive slow Ca(2+) oscillations in islets are data showing that oscillations in Ca(2+), mitochondrial oxygen consumption, and NAD(P)H levels are all terminated by membrane hyperpolarization. We demonstrate that these data are in fact compatible with a model in which glycolytic oscillations are the key player in rhythmic islet activity. Finally, we use the model to address the recent finding that the activity of islets from some mice is uniformly fast, whereas that from islets of other mice is slow. We propose a mechanism for this dichotomy.

This model was taken from the [CellML repository](#) and automatically converted to SBML. The original model was: [Bertram, Satin, Pedersen, Luciani, Sherman, 2007 version 02](#)

The original CellML model was created and curated by:

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

## 2 Unit Definitions

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

### 2.1 Unit substance

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

**Definition** mol

## 2.2 Unit volume

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

**Definition** l

## 2.3 Unit area

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

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

## 2.4 Unit length

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

**Definition** m

## 2.5 Unit time

**Notes** Second is the predefined SBML unit for time.

**Definition** s

# 3 Compartment

This model contains one compartment.

Table 2: Properties of all compartments.

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

### 3.1 Compartment Compartment

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

**Name** Compartment

## 4 Species

This model contains eleven species. Section 7 provides further details and the derived rates of change of each species.

Table 3: Properties of each species.

Id	Name	Compartment	Derived Unit	Constant	Boundary Condi- tion
Vm	Vm	Compartment	$\text{mol} \cdot \text{l}^{-1}$	$\square$	$\square$
n	n	Compartment	$\text{mol} \cdot \text{l}^{-1}$	$\square$	$\square$
G6P	G6P	Compartment	$\text{mol} \cdot \text{l}^{-1}$	$\square$	$\square$
FBP	FBP	Compartment	$\text{mol} \cdot \text{l}^{-1}$	$\square$	$\square$
NADHm	NADHm	Compartment	$\text{mol} \cdot \text{l}^{-1}$	$\square$	$\square$
delta_psi	delta_psi	Compartment	$\text{mol} \cdot \text{l}^{-1}$	$\square$	$\square$
Cam	Cam	Compartment	$\text{mol} \cdot \text{l}^{-1}$	$\square$	$\square$
ADPm	ADPm	Compartment	$\text{mol} \cdot \text{l}^{-1}$	$\square$	$\square$
adp	adp	Compartment	$\text{mol} \cdot \text{l}^{-1}$	$\square$	$\square$
c	c	Compartment	$\text{mol} \cdot \text{l}^{-1}$	$\square$	$\square$
Caer	Caer	Compartment	$\text{mol} \cdot \text{l}^{-1}$	$\square$	$\square$

## 5 Parameters

This model contains 151 global parameters.

Table 4: Properties of each parameter.

Id	Name	SBO	Value	Unit	Constant
cm	cm	0000258	5300.000		<input checked="" type="checkbox"/>
Ik	Ik		0.000		<input type="checkbox"/>
gK	gK	0000009	2700.000		<input checked="" type="checkbox"/>
n_infinity	n_infinity		$1.50710358059757 \cdot 10^{-4}$		<input type="checkbox"/>
tau_n	tau_n		20.000		<input checked="" type="checkbox"/>
Ica	Ica		-2927.842		<input type="checkbox"/>
gCa	gCa	0000009	1000.000		<input checked="" type="checkbox"/>
m_infinity	m_infinity		0.034		<input type="checkbox"/>
Ikca	Ikca		466.296		<input type="checkbox"/>
gkCa	gkCa	0000009	300.000		<input checked="" type="checkbox"/>
kd	kd	0000009	0.500		<input checked="" type="checkbox"/>
Ikatp	Ikatp		2433.430		<input type="checkbox"/>
gkATP_	gkATP_		16000.000		<input checked="" type="checkbox"/>
katpo	katpo		0.010		<input type="checkbox"/>
topo	topo		110.234		<input type="checkbox"/>
bottomo	bottomo		10871.926		<input type="checkbox"/>
mgadp	mgadp		187.605		<input type="checkbox"/>
adp3m	adp3m		153.495		<input type="checkbox"/>
atp4m	atp4m		68.150		<input type="checkbox"/>
JGPDH	JGPDH		$7.34846922834953 \cdot 10^{-4}$		<input type="checkbox"/>
kGPDH	kGPDH	0000009	$5 \cdot 10^{-4}$		<input checked="" type="checkbox"/>
F6P	F6P		90.300		<input type="checkbox"/>
JPFK	JPFK		0.374		<input type="checkbox"/>
JPFK_ms	JPFK_ms		$3.74364085279847 \cdot 10^{-4}$		<input type="checkbox"/>
bottom1	bottom1	0000009	1.000		<input checked="" type="checkbox"/>
weight1	weight1	0000009	1.000		<input checked="" type="checkbox"/>
topa1	topa1	0000009	0.000		<input checked="" type="checkbox"/>
k1	k1	0000009	30.000		<input checked="" type="checkbox"/>
k2	k2	0000009	1.000		<input checked="" type="checkbox"/>
k3	k3	0000009	50000.000		<input checked="" type="checkbox"/>
k4	k4	0000009	220.000		<input checked="" type="checkbox"/>
VmaxPFK	VmaxPFK	0000009	5.000		<input checked="" type="checkbox"/>
weight2	weight2		8444.405		<input type="checkbox"/>
topa2	topa2		0.000		<input type="checkbox"/>
bottom2	bottom2		8445.405		<input type="checkbox"/>
topa3	topa3		0.163		<input type="checkbox"/>
weight3	weight3		0.163		<input type="checkbox"/>

Id	Name	SBO	Value	Unit	Constant
bottom3	bottom3		8445.568		<input type="checkbox"/>
f13	f13	0000009	0.020		<input checked="" type="checkbox"/>
f43	f43	0000009	20.000		<input checked="" type="checkbox"/>
f23	f23	0000009	0.200		<input checked="" type="checkbox"/>
f42	f42	0000009	20.000		<input checked="" type="checkbox"/>
f41	f41	0000009	20.000		<input checked="" type="checkbox"/>
weight4	weight4		68.856		<input type="checkbox"/>
topa4	topa4		69.020		<input type="checkbox"/>
bottom4	bottom4		8514.424		<input type="checkbox"/>
weight5	weight5		2.160		<input type="checkbox"/>
topa5	topa5		69.020		<input type="checkbox"/>
bottom5	bottom5		8516.584		<input type="checkbox"/>
weight6	weight6		911.996		<input type="checkbox"/>
topa6	topa6		69.020		<input type="checkbox"/>
bottom6	bottom6		9428.580		<input type="checkbox"/>
weight7	weight7		1.761		<input type="checkbox"/>
topa7	topa7		70.781		<input type="checkbox"/>
bottom7	bottom7		9430.341		<input type="checkbox"/>
weight8	weight8		37.182		<input type="checkbox"/>
topa8	topa8		107.963		<input type="checkbox"/>
bottom8	bottom8		9467.524		<input type="checkbox"/>
weight9	weight9		16.667		<input type="checkbox"/>
topa9	topa9		107.963		<input type="checkbox"/>
bottom9	bottom9		9484.190		<input type="checkbox"/>
weight10	weight10		7037.004		<input type="checkbox"/>
topa10	topa10		107.963		<input type="checkbox"/>
bottom10	bottom10		16521.194		<input type="checkbox"/>
weight11	weight11		135.902		<input type="checkbox"/>
topa11	topa11		243.865		<input type="checkbox"/>
bottom11	bottom11		16657.095		<input type="checkbox"/>
weight12	weight12		2869.018		<input type="checkbox"/>
topa12	topa12		3112.883		<input type="checkbox"/>
bottom12	bottom12		19526.114		<input type="checkbox"/>
weight13	weight13		36.000		<input type="checkbox"/>
topa13	topa13		3112.883		<input type="checkbox"/>
bottom13	bottom13		19562.114		<input type="checkbox"/>
weight14	weight14		759.996		<input type="checkbox"/>
topa14	topa14		3112.883		<input type="checkbox"/>
bottom14	bottom14		20322.110		<input type="checkbox"/>
weight15	weight15		1467.736		<input type="checkbox"/>
topa15	topa15		3112.883		<input type="checkbox"/>
bottom15	bottom15		21789.846		<input type="checkbox"/>

Id	Name	SBO	Value	Unit	Constant
weight16	weight16		1549.270		<input type="checkbox"/>
topa16	topa16		4662.153		<input type="checkbox"/>
bottom16	bottom16		23339.116		<input type="checkbox"/>
topb	topb		1467.736		<input type="checkbox"/>
lambda	lambda	0000009	0.060		<input checked="" type="checkbox"/>
JPDH	JPDH	0000009	0.452		<input type="checkbox"/>
p1	p1	0000009	400.000		<input checked="" type="checkbox"/>
p2	p2	0000009	1.000		<input checked="" type="checkbox"/>
p3	p3	0000009	0.010		<input checked="" type="checkbox"/>
JGPDHbas	JGPDHbas	0000009	$5 \cdot 10^{-4}$		<input checked="" type="checkbox"/>
JO	JO		0.447		<input type="checkbox"/>
p4	p4	0000009	0.600		<input checked="" type="checkbox"/>
p5	p5	0000009	0.100		<input checked="" type="checkbox"/>
p6	p6	0000009	177.000		<input checked="" type="checkbox"/>
p7	p7	0000009	5.000		<input checked="" type="checkbox"/>
NADm	NADm		9.600		<input type="checkbox"/>
NADmtot	NADmtot	0000009	10.000		<input checked="" type="checkbox"/>
Cmito	Cmito	0000009	1.800		<input checked="" type="checkbox"/>
JHres	JHres		5.213		<input type="checkbox"/>
p8	p8	0000009	7.000		<input checked="" type="checkbox"/>
p9	p9		0.100		<input checked="" type="checkbox"/>
p10	p10	0000009	177.000		<input checked="" type="checkbox"/>
p11	p11	0000009	5.000		<input checked="" type="checkbox"/>
JF1F0	JF1F0		1.129		<input type="checkbox"/>
p13	p13		10.000		<input checked="" type="checkbox"/>
p14	p14		190.000		<input checked="" type="checkbox"/>
p15	p15	0000009	8.500		<input checked="" type="checkbox"/>
p16	p16	0000009	35.000		<input checked="" type="checkbox"/>
JHatp	JHatp		3.387		<input type="checkbox"/>
JGK_ms	JGK_ms		$4 \cdot 10^{-4}$		<input type="checkbox"/>
JGK	JGK	0000009	0.400		<input checked="" type="checkbox"/>
JHleak	JHleak		0.298		<input type="checkbox"/>
p17	p17	0000009	0.002		<input checked="" type="checkbox"/>
p18	p18	0000009	-0.030		<input checked="" type="checkbox"/>
JANT	JANT		1.124		<input type="checkbox"/>
p19	p19	0000009	0.350		<input checked="" type="checkbox"/>
p20	p20	0000009	2.000		<input checked="" type="checkbox"/>
FRT	FRT	0000009	0.037		<input checked="" type="checkbox"/>
RATm	RATm		0.351		<input type="checkbox"/>
Juni	Juni		0.158		<input type="checkbox"/>
p21	p21	0000009	0.040		<input checked="" type="checkbox"/>
p22	p22	0000009	1.100		<input checked="" type="checkbox"/>

Id	Name	SBO	Value	Unit	Constant
JNaCa	JNaCa		0.162		<input type="checkbox"/>
p23	p23	0000009	0.010		<input checked="" type="checkbox"/>
p24	p24	0000009	0.016		<input checked="" type="checkbox"/>
fmito	fmito	0000009	0.010		<input checked="" type="checkbox"/>
Jmito	Jmito		0.004		<input type="checkbox"/>
ATPm	ATPm		3.900		<input type="checkbox"/>
Amtot	Amtot	0000009	15.000		<input checked="" type="checkbox"/>
Jhyd	Jhyd		0.080		<input type="checkbox"/>
khyd	khyd	0000009	$5 \cdot 10^{-5}$		<input checked="" type="checkbox"/>
khydbas	khydbas	0000009	$5 \cdot 10^{-5}$		<input checked="" type="checkbox"/>
atp	atp		1363.000		<input type="checkbox"/>
atot	atot	0000009	2500.000		<input checked="" type="checkbox"/>
fcyt	fcyt	0000009	0.010		<input checked="" type="checkbox"/>
Jmem	Jmem		0.001		<input type="checkbox"/>
kPMCA	kPMCA	0000009	0.100		<input checked="" type="checkbox"/>
alpha	alpha	0000009	$4.5 \cdot 10^{-6}$		<input checked="" type="checkbox"/>
Cbas	Cbas	0000009	0.050		<input checked="" type="checkbox"/>
Jleak	Jleak		0.069		<input type="checkbox"/>
pleak	pleak	0000009	$2 \cdot 10^{-4}$		<input checked="" type="checkbox"/>
JSERCA	JSERCA		0.068		<input type="checkbox"/>
kSERCA	kSERCA	0000009	0.400		<input checked="" type="checkbox"/>
Jer	Jer		$9.659999999999995 \cdot 10^{-4}$		<input type="checkbox"/>
fer	fer	0000009	0.010		<input checked="" type="checkbox"/>
Vc_Ver	Vc_Ver	0000009	31.000		<input checked="" type="checkbox"/>
gamma	gamma	0000009	0.001		<input checked="" type="checkbox"/>
kappa	kappa	0000009	0.001		<input checked="" type="checkbox"/>
delta	delta		0.073		<input type="checkbox"/>
VK	VK	0000009	-75.000		<input checked="" type="checkbox"/>
VCa	VCa	0000009	25.000		<input checked="" type="checkbox"/>
AMP	AMP	0000009	500.000		<input checked="" type="checkbox"/>

## 6 Rules

This is an overview of 94 rules.

### 6.1 Rule $I_k$

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

$$I_k = g_K \cdot [n] \cdot ([V_m] - V_K) \quad (1)$$



## 6.2 Rule `n_infinity`

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

$$n\_infinity = \frac{1}{1 + \exp\left(\frac{(16+[Vm])}{5}\right)} \quad (2)$$

## 6.3 Rule `m_infinity`

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

$$m\_infinity = \frac{1}{1 + \exp\left(\frac{(20+[Vm])}{12}\right)} \quad (3)$$

## 6.4 Rule `Ica`

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

$$Ica = gCa \cdot m\_infinity \cdot ([Vm] - VCa) \quad (4)$$

## 6.5 Rule `Ikca`

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

$$Ikca = gkCa \cdot \frac{[c]^2}{[c]^2 + kd^2} \cdot ([Vm] - VK) \quad (5)$$

## 6.6 Rule `mgadp`

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

$$mgadp = 0.165 \cdot [adp] \quad (6)$$

## 6.7 Rule `topo`

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

$$topo = 0.08 \cdot \left(1 + \frac{2 \cdot mgadp}{17}\right) + 0.89 \cdot \left(\frac{mgadp}{17}\right)^2 \quad (7)$$

## 6.8 Rule `adp3m`

Rule `adp3m` is an assignment rule for parameter `adp3m`:

$$adp3m = 0.135 \cdot [adp] \quad (8)$$

### 6.9 Rule JGPDH

Rule JGPDH is an assignment rule for parameter JGPDH:

$$\text{JGPDH} = \text{kGPDH} \cdot [\text{FBP}]^{\frac{1}{2}} \quad (9)$$

### 6.10 Rule F6P

Rule F6P is an assignment rule for parameter F6P:

$$\text{F6P} = 0.3 \cdot [\text{G6P}] \quad (10)$$

### 6.11 Rule topa2

Rule topa2 is an assignment rule for parameter topa2:

$$\text{topa2} = \text{topa1} \quad (11)$$

### 6.12 Rule weight3

Rule weight3 is an assignment rule for parameter weight3:

$$\text{weight3} = \frac{\text{F6P}^2}{\text{k3}} \quad (12)$$

### 6.13 Rule topa3

Rule topa3 is an assignment rule for parameter topa3:

$$\text{topa3} = \text{topa2} + \text{weight3} \quad (13)$$

### 6.14 Rule weight5

Rule weight5 is an assignment rule for parameter weight5:

$$\text{weight5} = \frac{[\text{FBP}]}{\text{k2}} \quad (14)$$

### 6.15 Rule weight7

Rule weight7 is an assignment rule for parameter weight7:

$$\text{weight7} = \frac{[\text{FBP}] \cdot \text{F6P}^2}{\text{k2} \cdot \text{f23} \cdot \text{k3}} \quad (15)$$

### 6.16 Rule weight9

Rule weight9 is an assignment rule for parameter weight9:

$$\text{weight9} = \frac{\text{AMP}}{\text{k1}} \quad (16)$$

### 6.17 Rule `weight11`

Rule `weight11` is an assignment rule for parameter `weight11`:

$$\text{weight11} = \frac{\text{AMP} \cdot \text{F6P}^2}{k1 \cdot k3 \cdot f13} \quad (17)$$

### 6.18 Rule `weight13`

Rule `weight13` is an assignment rule for parameter `weight13`:

$$\text{weight13} = \frac{\text{AMP} \cdot [\text{FBP}]}{k1 \cdot k2} \quad (18)$$

### 6.19 Rule `weight15`

Rule `weight15` is an assignment rule for parameter `weight15`:

$$\text{weight15} = \frac{\text{AMP} \cdot [\text{FBP}] \cdot \text{F6P}^2}{k1 \cdot k2 \cdot f23 \cdot f13 \cdot k3} \quad (19)$$

### 6.20 Rule `topb`

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

$$\text{topb} = \text{weight15} \quad (20)$$

### 6.21 Rule `J0`

Rule `J0` is an assignment rule for parameter `J0`:

$$J0 = \frac{\frac{p4 \cdot [\text{NADHm}]}{p5 + [\text{NADHm}]}}{1 + \exp\left(\frac{[\text{delta\_psi}] - p6}{p7}\right)} \quad (21)$$

### 6.22 Rule `NADm`

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

$$\text{NADm} = \text{NADmtot} - [\text{NADHm}] \quad (22)$$

### 6.23 Rule `JPDH`

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

$$JPDH = \frac{p1}{p2 + \frac{[\text{NADHm}]}{\text{NADm}}} \cdot \frac{[\text{Cam}]}{p3 + [\text{Cam}]} \cdot (\text{JGPDH} + \text{JGPDHbas}) \quad (23)$$

### 6.24 Rule JHres

Rule JHres is an assignment rule for parameter JHres:

$$\text{JHres} = p8 \cdot \frac{[\text{NADHm}]}{p9 + [\text{NADHm}]} \cdot \frac{1}{1 + \exp\left(\frac{[\text{delta\_psi}] - p10}{p11}\right)} \quad (24)$$

### 6.25 Rule JGK\_ms

Rule JGK\_ms is an assignment rule for parameter JGK\_ms:

$$\text{JGK\_ms} = \text{kappa} \cdot \text{JGK} \quad (25)$$

### 6.26 Rule JHleak

Rule JHleak is an assignment rule for parameter JHleak:

$$\text{JHleak} = p17 \cdot [\text{delta\_psi}] + p18 \quad (26)$$

### 6.27 Rule Juni

Rule Juni is an assignment rule for parameter Juni:

$$\text{Juni} = (p21 \cdot [\text{delta\_psi}] - p22) \cdot [c]^2 \quad (27)$$

### 6.28 Rule JNaCa

Rule JNaCa is an assignment rule for parameter JNaCa:

$$\text{JNaCa} = p23 \cdot \frac{[\text{Cam}]}{[c]} \cdot \exp(p24 \cdot [\text{delta\_psi}]) \quad (28)$$

### 6.29 Rule Jmito

Rule Jmito is an assignment rule for parameter Jmito:

$$\text{Jmito} = \text{JNaCa} - \text{Juni} \quad (29)$$

### 6.30 Rule ATPm

Rule ATPm is an assignment rule for parameter ATPm:

$$\text{ATPm} = \text{Amtot} - [\text{ADPm}] \quad (30)$$

### 6.31 Rule JF1F0

Rule JF1F0 is an assignment rule for parameter JF1F0:

$$\text{JF1F0} = \frac{p16 \cdot p13}{p13 + \text{ATPm}} \cdot \frac{1}{1 + \exp\left(\frac{p14 - [\text{delta\_psi}]}{p15}\right)} \quad (31)$$

### 6.32 Rule JHatp

Rule JHatp is an assignment rule for parameter JHatp:

$$\text{JHatp} = 3 \cdot \text{JF1F0} \quad (32)$$

### 6.33 Rule RATm

Rule RATm is an assignment rule for parameter RATm:

$$\text{RATm} = \frac{\text{ATPm}}{[\text{ADPm}]} \quad (33)$$

### 6.34 Rule JANT

Rule JANT is an assignment rule for parameter JANT:

$$\text{JANT} = \text{p19} \cdot \frac{\text{RATm}}{\text{RATm} + \text{p20}} \cdot \exp(0.5 \cdot \text{FRT} \cdot [\text{delta\_psi}]) \quad (34)$$

### 6.35 Rule atp

Rule atp is an assignment rule for parameter atp:

$$\text{atp} = \text{atot} - [\text{adp}] \quad (35)$$

### 6.36 Rule atp4m

Rule atp4m is an assignment rule for parameter atp4m:

$$\text{atp4m} = 0.05 \cdot \text{atp} \quad (36)$$

### 6.37 Rule bottomo

Rule bottomo is an assignment rule for parameter bottomo:

$$\text{bottomo} = \left(1 + \frac{\text{mgadp}}{17}\right)^2 \cdot \left(1 + \frac{\text{adp3m}}{26} + \frac{\text{atp4m}}{1}\right) \quad (37)$$

### 6.38 Rule katpo

Rule katpo is an assignment rule for parameter katpo:

$$\text{katpo} = \frac{\text{topo}}{\text{bottomo}} \quad (38)$$

### 6.39 Rule Ikatp

Rule Ikatp is an assignment rule for parameter Ikatp:

$$\text{Ikatp} = \text{gkATP}_- \cdot \text{katpo} \cdot ([\text{Vm}] - \text{VK}) \quad (39)$$

#### 6.40 Rule `weight2`

Rule `weight2` is an assignment rule for parameter `weight2`:

$$\text{weight2} = \frac{\text{atp}^2}{k4} \quad (40)$$

#### 6.41 Rule `bottom2`

Rule `bottom2` is an assignment rule for parameter `bottom2`:

$$\text{bottom2} = \text{bottom1} + \text{weight2} \quad (41)$$

#### 6.42 Rule `bottom3`

Rule `bottom3` is an assignment rule for parameter `bottom3`:

$$\text{bottom3} = \text{bottom2} + \text{weight3} \quad (42)$$

#### 6.43 Rule `weight4`

Rule `weight4` is an assignment rule for parameter `weight4`:

$$\text{weight4} = \frac{(\text{F6P} \cdot \text{atp})^2}{k3 \cdot k4 \cdot f43} \quad (43)$$

#### 6.44 Rule `topa4`

Rule `topa4` is an assignment rule for parameter `topa4`:

$$\text{topa4} = \text{topa3} + \text{weight4} \quad (44)$$

#### 6.45 Rule `bottom4`

Rule `bottom4` is an assignment rule for parameter `bottom4`:

$$\text{bottom4} = \text{bottom3} + \text{weight4} \quad (45)$$

#### 6.46 Rule `topa5`

Rule `topa5` is an assignment rule for parameter `topa5`:

$$\text{topa5} = \text{topa4} \quad (46)$$

#### 6.47 Rule `bottom5`

Rule `bottom5` is an assignment rule for parameter `bottom5`:

$$\text{bottom5} = \text{bottom4} + \text{weight5} \quad (47)$$

#### 6.48 Rule `weight6`

Rule `weight6` is an assignment rule for parameter `weight6`:

$$\text{weight6} = \frac{[\text{FBP}] \cdot \text{atp}^2}{k2 \cdot f42 \cdot k4} \quad (48)$$

#### 6.49 Rule `topa6`

Rule `topa6` is an assignment rule for parameter `topa6`:

$$\text{topa6} = \text{topa5} \quad (49)$$

#### 6.50 Rule `bottom6`

Rule `bottom6` is an assignment rule for parameter `bottom6`:

$$\text{bottom6} = \text{bottom5} + \text{weight6} \quad (50)$$

#### 6.51 Rule `topa7`

Rule `topa7` is an assignment rule for parameter `topa7`:

$$\text{topa7} = \text{topa6} + \text{weight7} \quad (51)$$

#### 6.52 Rule `bottom7`

Rule `bottom7` is an assignment rule for parameter `bottom7`:

$$\text{bottom7} = \text{bottom6} + \text{weight7} \quad (52)$$

#### 6.53 Rule `weight8`

Rule `weight8` is an assignment rule for parameter `weight8`:

$$\text{weight8} = \frac{[\text{FBP}] \cdot \text{F6P}^2 \cdot \text{atp}^2}{k2 \cdot f23 \cdot f42 \cdot f43 \cdot k3 \cdot k4} \quad (53)$$

#### 6.54 Rule `topa8`

Rule `topa8` is an assignment rule for parameter `topa8`:

$$\text{topa8} = \text{topa7} + \text{weight8} \quad (54)$$

#### 6.55 Rule `topa9`

Rule `topa9` is an assignment rule for parameter `topa9`:

$$\text{topa9} = \text{topa8} \quad (55)$$

### 6.56 Rule `bottom8`

Rule `bottom8` is an assignment rule for parameter `bottom8`:

$$\text{bottom8} = \text{bottom7} + \text{weight8} \quad (56)$$

### 6.57 Rule `bottom9`

Rule `bottom9` is an assignment rule for parameter `bottom9`:

$$\text{bottom9} = \text{bottom8} + \text{weight9} \quad (57)$$

### 6.58 Rule `weight10`

Rule `weight10` is an assignment rule for parameter `weight10`:

$$\text{weight10} = \frac{\text{AMP} \cdot \text{atp}^2}{k1 \cdot k4 \cdot f41} \quad (58)$$

### 6.59 Rule `topa10`

Rule `topa10` is an assignment rule for parameter `topa10`:

$$\text{topa10} = \text{topa9} \quad (59)$$

### 6.60 Rule `bottom10`

Rule `bottom10` is an assignment rule for parameter `bottom10`:

$$\text{bottom10} = \text{bottom9} + \text{weight10} \quad (60)$$

### 6.61 Rule `topa11`

Rule `topa11` is an assignment rule for parameter `topa11`:

$$\text{topa11} = \text{topa10} + \text{weight11} \quad (61)$$

### 6.62 Rule `bottom11`

Rule `bottom11` is an assignment rule for parameter `bottom11`:

$$\text{bottom11} = \text{bottom10} + \text{weight11} \quad (62)$$

### 6.63 Rule `weight12`

Rule `weight12` is an assignment rule for parameter `weight12`:

$$\text{weight12} = \frac{\text{AMP} \cdot \text{F6P}^2 \cdot \text{atp}^2}{k1 \cdot f13 \cdot f41 \cdot f43 \cdot k3 \cdot k4} \quad (63)$$



#### 6.64 Rule `topa12`

Rule `topa12` is an assignment rule for parameter `topa12`:

$$\text{topa12} = \text{topa11} + \text{weight12} \quad (64)$$

#### 6.65 Rule `bottom12`

Rule `bottom12` is an assignment rule for parameter `bottom12`:

$$\text{bottom12} = \text{bottom11} + \text{weight12} \quad (65)$$

#### 6.66 Rule `topa13`

Rule `topa13` is an assignment rule for parameter `topa13`:

$$\text{topa13} = \text{topa12} \quad (66)$$

#### 6.67 Rule `bottom13`

Rule `bottom13` is an assignment rule for parameter `bottom13`:

$$\text{bottom13} = \text{bottom12} + \text{weight13} \quad (67)$$

#### 6.68 Rule `weight14`

Rule `weight14` is an assignment rule for parameter `weight14`:

$$\text{weight14} = \frac{\text{AMP} \cdot [\text{FBP}] \cdot \text{atp}^2}{k1 \cdot k2 \cdot f42 \cdot f41 \cdot k4} \quad (68)$$

#### 6.69 Rule `topa14`

Rule `topa14` is an assignment rule for parameter `topa14`:

$$\text{topa14} = \text{topa13} \quad (69)$$

#### 6.70 Rule `bottom14`

Rule `bottom14` is an assignment rule for parameter `bottom14`:

$$\text{bottom14} = \text{bottom13} + \text{weight14} \quad (70)$$

#### 6.71 Rule `topa15`

Rule `topa15` is an assignment rule for parameter `topa15`:

$$\text{topa15} = \text{topa14} \quad (71)$$

### 6.72 Rule `bottom15`

Rule `bottom15` is an assignment rule for parameter `bottom15`:

$$\text{bottom15} = \text{bottom14} + \text{weight15} \quad (72)$$

### 6.73 Rule `weight16`

Rule `weight16` is an assignment rule for parameter `weight16`:

$$\text{weight16} = \frac{\text{AMP} \cdot [\text{FBP}] \cdot \text{F6P}^2 \cdot \text{atp}^2}{k1 \cdot k2 \cdot k3 \cdot k4 \cdot f23 \cdot f13 \cdot f42 \cdot f41 \cdot f43} \quad (73)$$

### 6.74 Rule `topa16`

Rule `topa16` is an assignment rule for parameter `topa16`:

$$\text{topa16} = \text{topa15} + \text{weight16} \quad (74)$$

### 6.75 Rule `bottom16`

Rule `bottom16` is an assignment rule for parameter `bottom16`:

$$\text{bottom16} = \text{bottom15} + \text{weight16} \quad (75)$$

### 6.76 Rule `JPFK`

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

$$\text{JPFK} = \frac{\text{VmaxPFK} \cdot \text{lambda} \cdot \text{topa16} + \text{VmaxPFK} \cdot \text{topb}}{\text{bottom16}} \quad (76)$$

### 6.77 Rule `JPFK_ms`

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

$$\text{JPFK\_ms} = \text{kappa} \cdot \text{JPFK} \quad (77)$$

### 6.78 Rule `Jhyd`

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

$$\text{Jhyd} = (\text{khyd} \cdot [\text{c}] + \text{khydbas}) \cdot \text{atp} \quad (78)$$

### 6.79 Rule `Jmem`

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

$$\text{Jmem} = (\text{alpha} \cdot \text{Ica} + \text{kPMCA} \cdot ([\text{c}] - \text{Cbas})) \quad (79)$$

### 6.80 Rule Jleak

Rule Jleak is an assignment rule for parameter Jleak:

$$Jleak = pleak \cdot ([Caer] - [c]) \quad (80)$$

### 6.81 Rule JSERCA

Rule JSERCA is an assignment rule for parameter JSERCA:

$$JSERCA = kSERCA \cdot [c] \quad (81)$$

### 6.82 Rule Jer

Rule Jer is an assignment rule for parameter Jer:

$$Jer = Jleak - JSERCA \quad (82)$$

### 6.83 Rule delta

Rule delta is an assignment rule for parameter delta:

$$delta = \frac{3.9}{53.2} \quad (83)$$

### 6.84 Rule Vm

Rule Vm is a rate rule for species Vm:

$$\frac{d}{dt} Vm = \frac{(Ik + Ica + Ikca + Ikatp)}{cm} \quad (84)$$

### 6.85 Rule n

Rule n is a rate rule for species n:

$$\frac{d}{dt} n = \frac{n\_infinity - [n]}{tau\_n} \quad (85)$$

### 6.86 Rule G6P

Rule G6P is a rate rule for species G6P:

$$\frac{d}{dt} G6P = JGK\_ms - JPFK\_ms \quad (86)$$

### 6.87 Rule FBP

Rule FBP is a rate rule for species FBP:

$$\frac{d}{dt} FBP = JPFK\_ms - 0.5 \cdot JGPDH \quad (87)$$

### 6.88 Rule NADHm

Rule NADHm is a rate rule for species NADHm:

$$\frac{d}{dt} \text{NADHm} = \text{gamma} \cdot (\text{JPDH} - \text{JO}) \quad (88)$$

### 6.89 Rule delta\_psi

Rule delta\_psi is a rate rule for species delta\_psi:

$$\frac{d}{dt} \text{delta\_psi} = \frac{\text{JHres} - (\text{JHatp} + \text{JANT} + \text{JHleak} + \text{JNaCa} + 2 \cdot \text{Juni})}{\text{Cmito}} \quad (89)$$

### 6.90 Rule Cam

Rule Cam is a rate rule for species Cam:

$$\frac{d}{dt} \text{Cam} = \text{fmito} \cdot \text{Jmito} \quad (90)$$

### 6.91 Rule ADPm

Rule ADPm is a rate rule for species ADPm:

$$\frac{d}{dt} \text{ADPm} = \text{gamma} \cdot (\text{JANT} - \text{JF1F0}) \quad (91)$$

### 6.92 Rule adp

Rule adp is a rate rule for species adp:

$$\frac{d}{dt} \text{adp} = \text{delta} \cdot \text{JANT} + \text{Jhyd} \quad (92)$$

### 6.93 Rule c

Rule c is a rate rule for species c:

$$\frac{d}{dt} \text{c} = \text{fcyt} \cdot (\text{Jmem} + \text{Jer} + \text{delta} \cdot \text{Jmito}) \quad (93)$$

### 6.94 Rule Caer

Rule Caer is a rate rule for species Caer:

$$\frac{d}{dt} \text{Caer} = \text{fer} \cdot \text{Vc\_Ver} \cdot \text{Jer} \quad (94)$$

## 7 Derived Rate Equations

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

### 7.1 Species $V_m$

**Name**  $V_m$

**Initial amount**  $-60$  mol

**Involved in rule**  $V_m$

One rule which determines this species' quantity.

### 7.2 Species $n$

**Name**  $n$

**Initial amount**  $0$  mol

**Involved in rule**  $n$

One rule which determines this species' quantity.

### 7.3 Species $G6P$

**Name**  $G6P$

**Initial amount**  $301$  mol

**Involved in rule**  $G6P$

One rule which determines this species' quantity.

### 7.4 Species $FBP$

**Name**  $FBP$

**Initial amount**  $2.16$  mol

**Involved in rule**  $FBP$

One rule which determines this species' quantity.

### 7.5 Species [NADHm](#)

**Name** NADHm

**Initial amount** 0.4 mol

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

One rule which determines this species' quantity.

### 7.6 Species [delta\\_psi](#)

**Name** delta\_psi

**SBO:0000009** kinetic constant

**Initial amount** 164 mol

**Involved in rule** [delta\\_psi](#)

One rule which determines this species' quantity.

### 7.7 Species [Cam](#)

**Name** Cam

**Initial amount** 0.2 mol

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

One rule which determines this species' quantity.

### 7.8 Species [ADPm](#)

**Name** ADPm

**Initial amount** 11.1 mol

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

One rule which determines this species' quantity.

### 7.9 Species [adp](#)

**Name** adp

**Initial amount** 1137 mol

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

One rule which determines this species' quantity.

## 7.10 Species <sup>c</sup>

**Name** <sup>c</sup>

**Initial amount** 0.17 mol

**Involved in rule** <sup>c</sup>

One rule which determines this species' quantity.

## 7.11 Species <sup>Caer</sup>

**Name** <sup>Caer</sup>

**Initial amount** 345 mol

**Involved in rule** <sup>Caer</sup>

One rule which determines this species' quantity.

# A Glossary of Systems Biology Ontology Terms

**SBO:0000009 kinetic constant:** Numerical parameter that quantifies the velocity of a chemical reaction

**SBO:0000258 capacitance:** Measure of the amount of electric charge stored (or separated) for a given electric potential. The unit of capacitance is the Farad

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