# **SBML Model Report**

# Model name: "Bertram2004\_PancreaticBetaCell\_modelB"



May 6, 2016

# 1 General Overview

This is a document in SBML Level 2 Version 4 format. This model was created by the following two authors: Ishan Ajmera<sup>1</sup> and Catherine Lloyd<sup>2</sup> at September 29<sup>th</sup> 2011 at 10:04 p.m. and last time modified at April eighth 2016 at 5:06 p.m. Table 1 gives an overview of the quantities of all components of this model.

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

Element	Quantity	Element	Quantity
compartment types	0	compartments	1
species types	0	species	7
events	0	constraints	0
reactions	0	function definitions	0
global parameters	113	unit definitions	0
rules	78	initial assignments	0

#### **Model Notes**

This a model from the article:

Calcium and glycolysis mediate multiple bursting modes in pancreatic islets.

Bertram R, Satin L, Zhang M, Smolen P, Sherman A. <u>Biophys J</u>2004 Nov;87(5):3074-87 15347584, **Abstract:** 

<sup>&</sup>lt;sup>1</sup>EMBL-EBI, ajmera@ebi.ac.uk

 $<sup>^2</sup> The \ University \ of \ Auckland, The \ Bioengineering \ Institute, \verb|c.lloyd@aukland.ac.nz| \\$ 

Pancreatic islets of Langerhans produce bursts of electrical activity whenexposed to stimulatory glucose levels. These bursts often have a regularrepeating pattern, with a period of 10-60 s. In some cases, however, the burstsare episodic, clustered into bursts of bursts, which we call compound bursting. Consistent with this are recordings of free Ca2+ concentration, oxygenconsumption, mitochondrial membrane potential, and intraislet glucose levels that exhibit very slow oscillations, with faster oscillations superimposed. Wedescribe a new mathematical model of the pancreatic beta-cell that can account for these multimodal patterns. The model includes the feedback of cytosolic Ca2+onto ion channels that can account for bursting, and a metabolic subsystem thatis capable of producing slow oscillations driven by oscillations in glycolysis. This slow rhythm is responsible for the slow mode of compound bursting in themodel. We also show that it is possible for glycolytic oscillations alone todrive a very slow form of bursting, which we call "glycolytic bursting., Finally, the model predicts that there is bistability between stationary andoscillatory glycolysis for a range of parameter values. We provide experimental support for this model prediction. Overall, the model can account for adiversity of islet behaviors described in the literature over the past 20 years.

This model was taken from the CellML repository and automatically converted to SBML. The original model was:Bertram R, Satin L, Zhang M, Smolen P, Sherman A. (2004) - version=1.0

The original CellML model was created by:

#### **Catherine Lloyd**

c.lloyd@auckland.ac.nz

The University of Auckland

This model originates from BioModels Database: A Database of Annotated Published Models (http://www.ebi.ac.uk/biomodels/). It is copyright (c) 2005-2011 The BioModels.net Team. For more information see the terms of use.

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** 1

# 2.3 Unit area

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

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

# 2.4 Unit length

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

**Definition** m

# 2.5 Unit time

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		

# 3.1 Compartment COMpartment

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

Name COMpartment

# 4 Species

This model contains seven 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
V	V	COMpartment	$\text{mol} \cdot l^{-1}$		
n	n	COMpartment	$\text{mol} \cdot 1^{-1}$		
С	c	COMpartment	$\text{mol} \cdot 1^{-1}$		
cer	cer	COMpartment	$\text{mol} \cdot l^{-1}$		
g6p	g6p	COMpartment	$\text{mol} \cdot l^{-1}$		$\Box$
fbp	fbp	COMpartment	$\text{mol} \cdot 1^{-1}$	$\Box$	
adp	adp	COMpartment	$\text{mol} \cdot 1^{-1}$		

# **5 Parameters**

This model contains 113 global parameters.

Table 4: Properties of each parameter.

Id	Name	SBO	Value	Unit	Constant
IK	IK		1012.500		
ICa	ICa		-2927.842		
IKCa	IKCa		1800.000		
Cm	Cm	0000258	5300.000		$\mathbf{Z}$
gK	gK	0000009	2700.000		$   \overline{\checkmark} $
gKCa	gKCa	0000009	600.000		$   \overline{\checkmark} $
kd	kd	0000009	0.500		
gCa	gCa	0000009	1000.000		$\overline{\mathbf{Z}}$
minf	minf		0.034		
VCa	VCa	0000009	25.000		
taun	taun		20.000		
ninf	ninf	1.5	50710358059757 · 10	$)^{-4}$	$\Box$
fcyt	fcyt	0000009	0.010		
Jmem	Jmem		-0.037		
Jer	Jer		-0.063		$\Box$
fer	fer	0000009	0.010		
${ t sigmaV}$	sigmaV	0000009	31.000		$   \overline{\mathbf{Z}} $
pleak	pleak	0000009	2 · 10	$)^{-4}$	
Kserca	Kserca	0000009	0.400		
lambdaer	lambdaer		1.000		
epser	epser	0000009	1.000		
alpha	alpha	0000009	4.5 · 10	$)^{-6}$	
kpmca	kpmca	0000009	0.200		
Jserca	Jserca		0.100		
Jleak	Jleak		0.037		
rgpdh	rgpdh		1.265		
Rgk	Rgk	0000009	0.200		
atot	atot		3000.000		
pfkbas	pfkbas	0000009	0.060		
f6p	f6p		60.000		
lambda	lambda	0000009	0.005		
pfk	pfk		0.551		
bottom1	bottom1		1.000		
topa1	topa1	0000009	0.000		
k1	k1	0000009	30.000		$   \overline{\checkmark} $
k2	k2	0000009	1.000		$ \overline{\checkmark} $
k3	k3	0000009	50000.000		$\square$

Id	Name	SBO	Value	Unit	Constant
k4	k4	0000009	1000.000		<b>✓</b>
cat	cat	0000009	2.000		$\overline{\mathbf{Z}}$
weight2	weight2		3609.038		
topa2	topa2		0.000		
bottom2	bottom2		3610.038		
topa3	topa3		0.072		
weight3	weight3		0.072		
bottom3	bottom3		3610.110		
famp	famp	0000009	0.020		$\square$
fatp	fatp	0000009	20.000		$\overline{\mathbf{Z}}$
ffbp	ffbp	0000009	0.200		$\overline{\mathbf{Z}}$
fbt	fbt	0000009	20.000		$\overline{\mathbf{Z}}$
fmt	fmt	0000009	20.000		$\overline{\mathbf{Z}}$
weight4	weight4		12.993		
topa4	topa4		13.065		
bottom4	bottom4		3623.102		
weight5	weight5		40.000		
topa5	topa5		13.065		
bottom5	bottom5		3663.102		
weight6	weight6		7218.076		
topa6	topa6		13.065		
bottom6	bottom6		10881.178		
weight7	weight7		14.400		
topa7	topa7		27.465		
bottom7	bottom7		10895.578		
weight8	weight8		129.925		
topa8	topa8		157.390		
bottom8	bottom8		11025.504		
weight9	weight9		10.675		
topa9	topa9		157.390		
bottom9	bottom9		11036.179		
weight10	weight10		1926.343		
topa10	topa10		157.390		
bottom10	bottom10		12962.522		
weight11	weight11		38.430		
topa11	topa11		195.820		
bottom11	bottom11		13000.952		
weight12	weight12		346.742		
topa12	topa12		542.562		
bottom12	bottom12		13347.694		
weight13	weight13		427.004		
topa13	topa13		542.562		

bottom13   bottom13   13774.698   □   weight14   weight14   3852.686   □   topa14   topa14   topa14   542.562   □   weight15   weight15   7686.077   □   topa15   topa15   542.562   □   bottom15   bottom15   25313.462   □   bottom16   weight16   weight16   weight16   weight16   weight16   topa16   topa16   topa16   topa16   bottom16   bottom16   28780.880   □   bottom6   bottom16   28780.880   □   mgadp   128.700   □   mgadp   mgadp   128.700   □   mgadp   adp3m   adp3m   adp3m   adp3m   atp4m   94.987   □   □   bottom0   bottom0   52.301   □   □   bottom0   bottom0   7348.241   □   □   katpo   katpo   0.007   □   EKATP   KKATP   2669.036   □   EKATP   KKATP   2669.036   □   EKATP   Kkdd   kdd   17.000   ✓   kdd   kdd   17.000   ✓   kdd   kdd   kdd   17.000   ✓   kdd   kdd   17.000   ✓   kdd   kdd   17.000   ✓   kdd   kdd   17.000   ✓   matpa   matpa   1899.747   □   EADS   matpa   mga   mga	Id	Name	SBO	Value	Unit	Constant
weight14         weight14         3852.686         □           topa14         topa14         542.562         □           bottom14         bottom15         □           weight15         weight15         7686.077         □           topa15         topa15         542.562         □           bottom15         bottom15         25313.462         □           weight16         weight16         3467.418         □           topa16         topa16         4009.980         □           bottom16         28780.880         □         □           topb         7686.077         □         □           mgadp         128.700         □         □           adp3m         adp3m         105.300         □           adp3m         adp3m         105.300         □           adp4m         49.4987         □         □           topo         52.301         □         □           bottomo         bottomo         7348.241         □         □           katpo         0.007         □         □         □           yK         VK         0000009         25000.000         □         □	bottom13	bottom13		13774.698		
topa14 bottom14 bottom14 17627.385 ⊟  weight15 weight15 7686.077 ⊟  topa15 topa15 542.562 ⊞  bottom15 bottom15 25313.462 ⊞  weight16 weight16 3467.418 ⊞  topa16 topa16 4009.980 ⊞  bottom16 bottom16 28780.880 ⊞  topb topb 7686.077 ⊞  mgadp mgadp 128.700 ⊞  adp3m adp3m 105.300 ⊞  adp4m atp4m 94.987 ⊞  topo topo 52.3301 ⊞  bottom0 bottom0 7348.241 ⊞  katpo katpo 0.007 ⊞  IKATP 1669.036 ⊞  VK VK 0000009 −75.000 ∰  gkatpbar gkatpbar 0000009 25000.000 ∰  kt kt kt 0000009 1.000 ∰  kt kt kt 0000009 1.000 ∰  r r r 0000009 1.000 ∰  kg kg 000009 1.000 ∰  r r r 0000009 1.000 ∰  r r r 0000009 1.000 ∰  r r r 0000009 1.000 ∰  kg kg kg 000009 1.000 ∰  kg kg kg 000009 1.000 ∰  rad pamp 320.253 ⊞  rad rad 1579.494 ⊞	weight14	weight14		3852.686		
bottom14         bottom14         17627.385         □           weight15         weight15         7686.077         □           topa15         topa15         542.562         □           bottom15         bottom15         25313.462         □           weight16         weight16         3467.418         □           topa16         topa16         4009.980         □           bottom16         28780.880         □           topb         topb         7686.077         □           mgadp         mgadp         128.700         □           adp3m         adp3m         105.300         □           atp4m         atp4.987         □         □           topo         52.301         □         □           bottomo         bottomo         7348.241         □         □           katpo         0.007         □         □         □           VK         VK         000009         -75.000         □         □           gkatpbar         gkatpbar         000009         25000.000         □         □           ktd         ktd         000009         260.00         □         □ <td< td=""><td>_</td><td>topa14</td><td></td><td>542.562</td><td></td><td></td></td<>	_	topa14		542.562		
weight15         topa15         542.562         ⊟           bottom15         bottom15         25313.462         ⊟           weight16         weight16         3467.418         ⊟           topa16         topa16         4009.980         ⊟           bottom16         bottom16         28780.880         ⊟           topb         topb         7686.077         ⊟           mgadp         128.700         ⊟           adp3m         alp3m         105.300         ⊟           atp4m         94.987         ⊟           topo         52.301         ⊟           bottomo         7348.241         ⊟           katpo         0.007         ⊟           KATP         2669.036         ⊟           VK         VK         000009         -75.000         ☑           gkatpbar         gkatpbar         000009         25000.000         ☑           ktd         ktd         000009         26.000         ☑           ktt         ktt         000009         1.000         ☑           atp         1899.747         ⊟         ⊟           taua         taua         1.247         ⊟      <	bottom14	bottom14		17627.385		
bottom15         bottom15         25313.462         □           weight16         weight16         3467.418         □           topa16         topa16         4009.980         □           bottom16         bottom16         28780.880         □           topb         topb         7686.077         □           mgadp         mgadp         128.700         □           adp3m         adp3m         105.300         □           atp4m         atp4m         94.987         □           topo         52.301         □         □           bottomo         bottomo         7348.241         □         □           katpo         0.007         □	weight15	weight15		7686.077		
bottom15         bottom15         25313.462         □           weight16         weigh16         3467.418         □           topa16         topa16         4009.980         □           bottom16         bottom16         28780.880         □           topb         topb         7686.077         □           mgadp         mgadp         128.700         □           adp3m         adp3m         105.300         □           atp4m         atp4m         94.987         □           topo         52.301         □         □           bottomo         bottomo         7348.241         □         □           katpo         0.007         □         □         □         □           VK         VK         000009         -75.000         □	topa15	topa15		542.562		
topa16 topa16 bottom16 28780.880 ⊟  topb topb 7686.077 ⊞  mgadp mgadp 128.700 ⊞  adp3m adp3m 105.300 ⊞  topo topo 52.301 ⊞  bottom0 bottom0 7348.241 ⊞  katpo katpo 0.0007 ⊞  IKATP IKATP 2669.036 ⊞  VK VK 000009 -75.000				25313.462		
bottom16 bottom16 28780.880 ⊟ topb topb 7686.077 ⊞ mgadp mgadp 128.700 ⊞ adp3m adp3m 105.300 ⊞ atp4m atp4m 94.987 topo topo 52.301 ⊞ bottomo bottomo 7348.241 ⊞ katpo katpo 0.007 ⊞ IKATP IKATP 2669.036 VK VK 0000009 −75.000 ☑ gkatpbar gkatpbar 0000009 25000.000 ☑ ktd ktd vd 000009 26.000 ☑ ktt ktt 0000009 1.000 ☑ atp atp 1899.747 ⊞ fback fback 1.247 taua taua 0000009 1.000 ☑ r1 r1 r1 0000009 0.350 r r r r 0000009 1.000 y y y 0.2447 vg kg kg 0000009 10.000 amp amp 320.253 rad 1579.494 ⊞	weight16	weight16		3467.418		
topb         109b         7686.077         □           mgadp         mgadp         128.700         □           adp3m         adp3m         105.300         □           atp4m         atp4m         94.987         □           topo         52.301         □           bottomo         bottomo         7348.241         □           katpo         0.007         □           KATP         2669.036         □           VK         VK         000009         25000.000           kdd         kdd         17.000         ✓           ktd         ktd         000009         26.000         ✓           ktt         ktt         0000009         26.000         ✓           ktt         ktt         0000009         1.000         ✓           atp         atp         1899.747         □         □           fback         fback         1.247         □         □           taua         taua         000009         0.350         ✓         ✓           r         r         0000009         1.000         ✓         ✓           yg         yg         2.200         ✓         <	topa16	topa16		4009.980		
mgadp         mgadp         128.700         □           adp3m         adp3m         105.300         □           atp4m         atp4m         94.987         □           topo         52.301         □           bottomo         bottomo         7348.241         □           katpo         0.0007         □           IKATP         1KATP         2669.036         □           VK         VK         000009         2-75.000         ☑           gkatpbar         gkatpbar         000009         25000.000         ☑           ktd         ktd         000009         26.000         ☑           ktt         ktt         000009         1.000         ☑           atp         atp         1899.747         □         □           fback         1.247         □         □           taua         taua         000009         0.350         ☑           r         r         0000009         1.000         ☑           yg         y         0.247         □           yg         y         0.247         □           yg         yg         2.200         ☑           <	bottom16	bottom16		28780.880		
mgadp         mgadp         128.700         □           adp3m         adp3m         105.300         □           atp4m         atp4m         94.987         □           topo         52.301         □           bottomo         bottomo         7348.241         □           katpo         0.0007         □           IKATP         1KATP         2669.036         □           VK         VK         000009         2-75.000         ☑           gkatpbar         gkatpbar         000009         25000.000         ☑           ktd         ktd         000009         26.000         ☑           ktt         ktt         000009         1.000         ☑           atp         atp         1899.747         □         □           fback         1.247         □         □           taua         taua         000009         0.350         ☑           r         r         0000009         1.000         ☑           yg         y         0.247         □           yg         y         0.247         □           yg         yg         2.200         ☑           <	topb	topb		7686.077		
adp3m       105.300       □         atp4m       34.987       □         topo       52.301       □         bottomo       7348.241       □         katpo       0.007       □         IKATP       2669.036       □         VK       VK       0000009       -75.000       ☑         gkatpbar       0000009       25000.000       ☑         kdd       kdd       17.000       ☑         ktd       ktd       0000009       26.000       ☑         ktt       kt       0000009       1.000       ☑         atp       atp       1899.747       □       □         fback       1.247       □       □         taua       taua       0000009       300000.000       ☑         r1       r1       0000009       0.350       ☑         r       r       0000009       1.000       ☑         yg       yg       2.200       ☑         kg       kg       0000009       10.000         amp       amp       320.253       □         rad       1579.494       □	mgadp	mgadp		128.700		
topo         52.301           bottomo         7348.241           katpo         0.007           IKATP         2669.036           VK         VK           VK         VK           gkatpbar         0000009           gkdd         ktd           ktd         17.000           ktd         ktd           ktt         0000009           atp         1.899.747           back         1.247           taua         taua           taua         1.247		adp3m		105.300		
topo         52.301         □           bottomo         7348.241         □           katpo         0.007         □           IKATP         IKATP         2669.036         □           VK         VK         0000009         -75.000         ☑           gkatpbar         gkatpbar         0000009         25000.000         ☑           ktd         ktd         17.000         ☑           ktd         ktd         0000009         26.000         ☑           ktt         ktt         0000009         1.000         ☑           atp         1899.747         □         □           fback         1.247         □         □           taua         taua         0000009         300000.000         ☑           r1         r1         0000009         0.350         ☑           r         r         0000009         1.000         ☑           yg         y         0.247         □           vg         kg         0000009         10.000           amp         320.253         □           rad         1579.494         □	atp4m	atp4m		94.987		
bottomo         5348.241         □           katpo         0.007         □           IKATP         2669.036         □           VK         VK         0000009         -75.000           gkatpbar         gkatpbar         0000009         25000.000           ktd         ktd         17.000         ✓           ktd         ktd         0000009         26.000         ✓           ktt         ktt         0000009         1.000         ✓           atp         1899.747         □         □           fback         1.247         □         □           taua         taua         0000009         300000.000         ✓           r1         r1         0000009         0.350         ✓           r         r         0000009         1.000         ✓           y         y         0.247         □           vg         y         2.200         ✓           kg         kg         0000009         10.000           amp         320.253         □           rad         1579.494         □				52.301		
katpo       0.007       ⊟         IKATP       2669.036       ⊟         VK       VK       0000009       -75.000       ✓         gkatpbar       gkatpbar       0000009       25000.000       ✓         ktd       ktd       17.000       ✓         ktt       ktd       0000009       26.000       ✓         ktt       ktt       0000009       1.000       ✓         atp       1899.747       ⊟       H         fback       fback       1.247       ⊟         taua       taua       0000009       300000.000       ✓         r1       r1       0000009       0.350       ✓         r       r       0000009       1.000       ✓         y       y       0.247       ⊟         vg       vg       2.200       ✓         kg       kg       000009       10.000       ✓         amp       320.253       ⊟         rad       1579.494       ⊟	bottomo			7348.241		
IKATP       2669.036       □         VK       VK       0000009       -75.000       ✓         gkatpbar       gkatpbar       0000009       25000.000       ✓         kdd       kdd       17.000       ✓         ktd       ktd       0000009       26.000       ✓         ktt       ktt       0000009       1.000       ✓         atp       1899.747       □       □         fback       1.247       □       □         taua       taua       0000009       300000.000       ✓         r1       r1       0000009       0.350       ✓         r       r       0000009       1.000       ✓         y       y       0.247       □         vg       vg       2.200       ✓         kg       kg       0000009       10.000       ✓         amp       amp       320.253       □         rad       1579.494       □	katpo	katpo		0.007		
gkatpbar       gkatpbar       0000009       25000.000         kdd       kdd       17.000         ktd       ktd       0000009       26.000         ktt       ktt       0000009       1.000         atp       1899.747       □         fback       fback       1.247         taua       taua       0000009       300000.000         r1       r1       0000009       0.350         r       r       0000009       1.000         y       y       0.247       □         vg       vg       2.200       ✓         kg       kg       0000009       10.000       ✓         amp       amp       320.253       □         rad       1579.494       □	IKATP	IKATP		2669.036		
gkatpbar       gkatpbar       0000009       25000.000       ✓         kdd       kdd       17.000       ✓         ktd       ktd       0000009       26.000         ktt       ktt       0000009       1.000         atp       1899.747       □         fback       1.247       □         taua       taua       0000009       300000.000         r1       r1       0000009       0.350       ✓         r       r       0000009       1.000       ✓         yg       yg       2.200       ✓         kg       kg       0000009       10.000       ✓         amp       320.253       □         rad       1579.494       □	VK	VK	0000009	-75.000		
kdd       ktd       17.000         ktd       ktd       0000009       26.000         ktt       ktt       0000009       1.000         atp       1899.747       □         fback       fback       1.247       □         taua       taua       0000009       300000.000       ✓         r1       r1       0000009       0.350       ✓         r       r       0000009       1.000       ✓         y       y       0.247       □         vg       vg       2.200       ✓         kg       kg       0000009       10.000       ✓         amp       amp       320.253       □         rad       1579.494       □	gkatpbar	gkatpbar	0000009	25000.000		
ktd       ktd       0000009       26.000         ktt       ktt       0000009       1.000         atp       atp       1899.747       □         fback       fback       1.247       □         taua       taua       0000009       300000.000       ☑         r1       r1       0000009       0.350       ☑         r       r       0000009       1.000       ☑         y       y       0.247       □         vg       vg       2.200       ☑         kg       kg       0000009       10.000       ☑         amp       amp       320.253       □         rad       1579.494       □	kdd	kdd		17.000		
ktt       ktt       0000009       1.000       ✓         atp       atp       1899.747       □         fback       fback       1.247       □         taua       taua       0000009       300000.000       ✓         r1       r1       0000009       0.350       ✓         r       r       0000009       1.000       ✓         yg       yg       2.200       ✓         kg       kg       0000009       10.000       ✓         amp       amp       320.253       □         rad       rad       1579.494       □	ktd	ktd	0000009	26.000		
atp       atp       1899.747       □         fback       1.247       □         taua       taua       0000009       300000.000         r1       r1       0000009       0.350         r       r       0000009       1.000         y       y       0.247         vg       vg       2.200         kg       kg       0000009       10.000         amp       320.253       □         rad       1579.494       □	ktt	ktt	0000009	1.000		
fback       1.247         taua       taua       0000009       300000.000         r1       r1       0000009       0.350         r       r       0000009       1.000         y       y       0.247         vg       vg       2.200         kg       kg       0000009       10.000         amp       320.253       □         rad       1579.494       □	atp	atp		1899.747		
r1       r1       0000009       0.350       ✓         r       r       0000009       1.000       ✓         y       y       0.247       □         vg       vg       2.200       ✓         kg       kg       0000009       10.000       ✓         amp       amp       320.253       □         rad       1579.494       □	fback	fback		1.247		
r1       r1       0000009       0.350       ✓         r       r       0000009       1.000       ✓         y       y       0.247       □         vg       vg       2.200       ✓         kg       kg       0000009       10.000       ✓         amp       amp       320.253       □         rad       1579.494       □	taua	taua	0000009	300000.000		
r       r       0000009       1.000       ✓         y       y       0.247       □         vg       vg       2.200       ✓         kg       kg       0000009       10.000       ✓         amp       amp       320.253       □         rad       1579.494       □	r1	r1	0000009	0.350		<u> </u>
y       y $0.247$ $\Box$ vg       vg $2.200$ $\Box$ kg       kg $0000009$ $10.000$ $\Box$ amp       amp $320.253$ $\Box$ rad $1579.494$ $\Box$	r	r	0000009	1.000		
vg $2.200$ kg       kg $0000009$ $10.000$ amp       amp $320.253$ $\Box$ rad $1579.494$ $\Box$	У	y		0.247		
kg kg 0000009 10.000 $\square$ amp amp 320.253 $\boxminus$ rad 1579.494	vg	vg		2.200		
amp amp $320.253$ $\Box$ rad $1579.494$ $\Box$			0000009	10.000		
	_			320.253		
	rad	rad		1579.494		
	ratio	ratio		2.436		

# 6 Rules

This is an overview of 78 rules.

# **6.1 Rule IK**

Rule IK is an assignment rule for parameter IK:

$$IK = gK \cdot [n] \cdot ([V] - VK) \tag{1}$$

#### 6.2 Rule IKCa

Rule IKCa is an assignment rule for parameter IKCa:

$$IKCa = \frac{gKCa}{1 + \left(\frac{kd}{[c]}\right)^2} \cdot ([V] - VK)$$
 (2)

#### 6.3 Rule minf

Rule minf is an assignment rule for parameter minf:

$$\min f = \frac{1}{1 + \exp\left(\frac{\left(20 + \frac{|V|}{1}\right)}{12}\right)}$$
 (3)

#### 6.4 Rule ICa

Rule ICa is an assignment rule for parameter ICa:

$$ICa = gCa \cdot minf \cdot ([V] - VCa)$$
(4)

#### 6.5 Rule ninf

Rule ninf is an assignment rule for parameter ninf:

$$ninf = \frac{1}{1 + exp\left(\frac{\left(16 + \frac{[V]}{I}\right)}{5}\right)}$$
 (5)

# 6.6 Rule Jmem

Rule Jmem is an assignment rule for parameter Jmem:

$$Jmem = (alpha \cdot ICa + kpmca \cdot [c])$$
 (6)

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

#### 6.7 Rule Jserca

Rule Jserca is an assignment rule for parameter Jserca:

$$Jserca = Kserca \cdot [c] \tag{7}$$

#### 6.8 Rule Jleak

Rule Jleak is an assignment rule for parameter Jleak:

$$Jleak = pleak \cdot ([cer] - [c])$$
(8)

#### 6.9 Rule Jer

Rule Jer is an assignment rule for parameter Jer:

$$Jer = \frac{epser \cdot (Jleak - Jserca)}{lambdaer}$$
 (9)

# 6.10 Rule rgpdh

Rule rgpdh is an assignment rule for parameter rgpdh:

$$rgpdh = 0.2 \cdot \left( \left| \frac{[fbp] \cdot 1}{1^2} \right| \right)^{\frac{1}{2}}$$
 (10)

# **6.11 Rule** f6p

Rule f6p is an assignment rule for parameter f6p:

$$f6p = 0.3 \cdot [g6p] \tag{11}$$

# 6.12 Rule topa2

Rule topa2 is an assignment rule for parameter topa2:

$$topa2 = topa1 (12)$$

# 6.13 Rule weight3

Rule weight3 is an assignment rule for parameter weight3:

$$weight3 = \frac{f6p^2}{k3 \cdot 1} \tag{13}$$

# 6.14 Rule topa3

Rule topa3 is an assignment rule for parameter topa3:

$$topa3 = topa2 + weight3 \tag{14}$$

# 6.15 Rule weight5

Rule weight5 is an assignment rule for parameter weight5:

$$weight5 = \frac{[fbp]}{k2}$$
 (15)

# 6.16 Rule weight7

Rule weight7 is an assignment rule for parameter weight7:

weight7 = 
$$\frac{[fbp] \cdot f6p^2}{k2 \cdot k3 \cdot ffbp \cdot 1}$$
 (16)

# 6.17 Rule mgadp

Rule mgadp is an assignment rule for parameter mgadp:

$$mgadp = 0.165 \cdot [adp] \tag{17}$$

# 6.18 Rule adp3m

Rule adp3m is an assignment rule for parameter adp3m:

$$adp3m = 0.135 \cdot [adp] \tag{18}$$

# 6.19 Rule topo

Rule topo is an assignment rule for parameter topo:

$$topo = 0.08 \cdot \left(1 + \frac{2 \cdot mgadp}{kdd \cdot 1}\right) + 0.89 \cdot \left(\frac{mgadp}{kdd \cdot 1}\right)^{2}$$
(19)

# **6.20** Rule y

Rule y is an assignment rule for parameter y:

$$y = vg \cdot \frac{rgpdh}{kg + rgpdh}$$
 (20)

# 6.21 Rule fback

Rule fback is an assignment rule for parameter fback:

$$fback = r + y \tag{21}$$

# 6.22 Rule rad

Rule rad is an assignment rule for parameter rad:

$$rad = \frac{\left(\left|\left(\left[adp\right] - atot\right)^2 - 4 \cdot \left[adp\right]^2\right|\right)^{\frac{1}{2}}}{1}$$
 (22)

# 6.23 Rule atp

Rule atp is an assignment rule for parameter atp:

$$atp = 0.5 \cdot (atot - [adp] + rad \cdot 1)$$
(23)

# 6.24 Rule weight2

Rule weight2 is an assignment rule for parameter weight2:

$$weight2 = \frac{atp^2}{k4 \cdot 1}$$
 (24)

#### 6.25 Rule bottom2

Rule bottom2 is an assignment rule for parameter bottom2:

$$bottom2 = bottom1 + weight2$$
 (25)

#### 6.26 Rule bottom3

Rule bottom3 is an assignment rule for parameter bottom3:

$$bottom3 = bottom2 + weight3$$
 (26)

# 6.27 Rule weight4

Rule weight4 is an assignment rule for parameter weight4:

weight4 = 
$$\frac{(f6p \cdot atp)^2}{fatp \cdot k3 \cdot k4 \cdot 1^2}$$
 (27)

# 6.28 Rule topa4

Rule topa4 is an assignment rule for parameter topa4:

$$topa4 = topa3 + weight4 (28)$$

# 6.29 Rule bottom4

Rule bottom4 is an assignment rule for parameter bottom4:

$$bottom4 = bottom3 + weight4$$
 (29)

# 6.30 Rule topa5

Rule topa5 is an assignment rule for parameter topa5:

$$topa5 = topa4 (30)$$

#### 6.31 Rule bottom5

Rule bottom5 is an assignment rule for parameter bottom5:

$$bottom5 = bottom4 + weight5$$
 (31)

# 6.32 Rule weight6

Rule weight6 is an assignment rule for parameter weight6:

weight6 = 
$$\frac{[fbp] \cdot atp^2}{k2 \cdot k4 \cdot fbt \cdot 1}$$
 (32)

# 6.33 Rule topa6

Rule topa6 is an assignment rule for parameter topa6:

$$topa6 = topa5 (33)$$

#### 6.34 Rule bottom6

Rule bottom6 is an assignment rule for parameter bottom6:

$$bottom6 = bottom5 + weight6$$
 (34)

# **6.35 Rule** topa7

Rule topa7 is an assignment rule for parameter topa7:

$$topa7 = topa6 + weight7 (35)$$

#### 6.36 Rule bottom7

Rule bottom7 is an assignment rule for parameter bottom7:

$$bottom7 = bottom6 + weight7$$
 (36)

# 6.37 Rule weight8

Rule weight8 is an assignment rule for parameter weight8:

weight8 = 
$$\frac{[fbp] \cdot f6p^2 \cdot atp^2}{k2 \cdot k3 \cdot k4 \cdot ffbp \cdot fbt \cdot fatp \cdot 1^2}$$
 (37)

#### 6.38 Rule topa8

Rule topa8 is an assignment rule for parameter topa8:

$$topa8 = topa7 + weight8 (38)$$

# 6.39 Rule topa9

Rule topa9 is an assignment rule for parameter topa9:

$$topa9 = topa8 \tag{39}$$

#### 6.40 Rule bottom8

Rule bottom8 is an assignment rule for parameter bottom8:

$$bottom8 = bottom7 + weight8 (40)$$

# **6.41 Rule** topa10

Rule topa10 is an assignment rule for parameter topa10:

$$topa10 = topa9 (41)$$

# 6.42 Rule atp4m

Rule atp4m is an assignment rule for parameter atp4m:

$$atp4m = 0.05 \cdot atp \tag{42}$$

#### 6.43 Rule bottomo

Rule bottomo is an assignment rule for parameter bottomo:

$$bottomo = \left(1 + \frac{mgadp}{kdd \cdot 1}\right)^{2} \cdot \left(1 + \frac{adp3m}{ktd \cdot 1} + \frac{atp4m}{ktt \cdot 1}\right) \tag{43}$$

# 6.44 Rule katpo

Rule katpo is an assignment rule for parameter katpo:

$$katpo = \frac{topo}{bottomo}$$
 (44)

# 6.45 Rule IKATP

Rule IKATP is an assignment rule for parameter IKATP:

$$IKATP = gkatpbar \cdot katpo \cdot ([V] - VK)$$
(45)

# 6.46 Rule amp

Rule amp is an assignment rule for parameter amp:

$$amp = \frac{[adp] \cdot [adp]}{atp} \tag{46}$$

# 6.47 Rule weight9

Rule weight9 is an assignment rule for parameter weight9:

$$weight9 = \frac{amp}{k1}$$
 (47)

#### 6.48 Rule bottom9

Rule bottom9 is an assignment rule for parameter bottom9:

$$bottom9 = bottom8 + weight9 (48)$$

# 6.49 Rule weight10

Rule weight10 is an assignment rule for parameter weight10:

$$weight10 = \frac{amp \cdot atp^2}{k1 \cdot k4 \cdot fmt \cdot 1}$$
(49)

#### 6.50 Rule bottom10

Rule bottom10 is an assignment rule for parameter bottom10:

$$bottom 10 = bottom 9 + weight 10 (50)$$

# 6.51 Rule weight11

Rule weight11 is an assignment rule for parameter weight11:

weight11 = 
$$\frac{\text{amp} \cdot \text{f6p}^2}{\text{k1} \cdot \text{k3} \cdot \text{famp} \cdot 1}$$
 (51)

# **6.52 Rule** topa11

Rule topa11 is an assignment rule for parameter topa11:

$$topa11 = topa10 + weight11 (52)$$

#### 6.53 Rule bottom11

Rule bottom11 is an assignment rule for parameter bottom11:

$$bottom11 = bottom10 + weight11$$
 (53)

# 6.54 Rule weight12

Rule weight12 is an assignment rule for parameter weight12:

weight12 = 
$$\frac{\text{amp} \cdot \text{f6p}^2 \cdot \text{atp}^2}{\text{k1} \cdot \text{k3} \cdot \text{k4} \cdot \text{famp} \cdot \text{fmt} \cdot \text{fatp} \cdot 1^2}$$
 (54)

# **6.55 Rule** topa12

Rule topa12 is an assignment rule for parameter topa12:

$$topa12 = topa11 + weight12$$
 (55)

#### 6.56 Rule bottom12

Rule bottom12 is an assignment rule for parameter bottom12:

$$bottom12 = bottom11 + weight12$$
 (56)

# 6.57 Rule weight13

Rule weight13 is an assignment rule for parameter weight13:

$$weight13 = \frac{amp \cdot [fbp]}{k1 \cdot k2}$$
 (57)

# **6.58 Rule** topa13

Rule topa13 is an assignment rule for parameter topa13:

$$topa13 = topa12 (58)$$

#### 6.59 Rule bottom13

Rule bottom13 is an assignment rule for parameter bottom13:

$$bottom 13 = bottom 12 + weight 13$$
 (59)

# 6.60 Rule weight14

Rule weight14 is an assignment rule for parameter weight14:

weight14 = 
$$\frac{\text{amp} \cdot [\text{fbp}] \cdot \text{atp}^2}{\text{k1} \cdot \text{k2} \cdot \text{k4} \cdot \text{fbt} \cdot \text{fmt} \cdot 1}$$
(60)

# **6.61 Rule** topa14

Rule topa14 is an assignment rule for parameter topa14:

$$topa14 = topa13 (61)$$

#### 6.62 Rule bottom14

Rule bottom14 is an assignment rule for parameter bottom14:

$$bottom 14 = bottom 13 + weight 14$$
 (62)

# 6.63 Rule weight15

Rule weight15 is an assignment rule for parameter weight15:

weight15 = 
$$\frac{\text{amp} \cdot [\text{fbp}] \cdot \text{f6p}^2}{\text{k1} \cdot \text{k2} \cdot \text{k3} \cdot \text{ffbp} \cdot \text{famp} \cdot 1}$$
(63)

# **6.64 Rule** topa15

Rule topa15 is an assignment rule for parameter topa15:

$$topa15 = topa14 (64)$$

#### 6.65 Rule bottom15

Rule bottom15 is an assignment rule for parameter bottom15:

$$bottom15 = bottom14 + weight15$$
 (65)

# 6.66 Rule weight16

Rule weight16 is an assignment rule for parameter weight16:

$$weight16 = \frac{amp \cdot [fbp] \cdot f6p^2 \cdot atp^2}{k1 \cdot k2 \cdot k3 \cdot k4 \cdot ffbp \cdot famp \cdot fbt \cdot fmt \cdot fatp \cdot 1^2}$$
 (66)

# **6.67 Rule** topa16

Rule topa16 is an assignment rule for parameter topa16:

$$topa16 = topa15 + weight16 \tag{67}$$

#### 6.68 Rule bottom16

Rule bottom16 is an assignment rule for parameter bottom16:

$$bottom16 = bottom15 + weight16 (68)$$

# 6.69 Rule topb

Rule topb is an assignment rule for parameter topb:

$$topb = weight15 (69)$$

# 6.70 Rule pfk

Rule pfk is an assignment rule for parameter pfk:

$$pfk = 1 \cdot \frac{pfkbas \cdot cat \cdot topa16 + cat \cdot topb}{bottom16}$$
(70)

#### 6.71 Rule ratio

Rule ratio is an assignment rule for parameter ratio:

$$ratio = \frac{atp}{[adp]} \tag{71}$$

# 6.72 Rule V

Rule V is a rate rule for species V:

$$\frac{\mathrm{d}}{\mathrm{d}t}V = \frac{(\mathrm{IK} + \mathrm{ICa} + \mathrm{IKCa} + \mathrm{IKATP})}{\mathrm{Cm}}$$
 (72)

# **6.73** Rule n

Rule n is a rate rule for species n:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathbf{n} = \frac{\mathrm{ninf} - [\mathbf{n}]}{\mathrm{taun}} \tag{73}$$

#### **6.74 Rule** c

Rule c is a rate rule for species c:

$$\frac{\mathrm{d}}{\mathrm{d}t}c = \mathrm{fcyt} \cdot (\mathrm{Jmem} + \mathrm{Jer}) \tag{74}$$

# **6.75 Rule** cer

Rule cer is a rate rule for species cer:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{cer} = \mathrm{fer} \cdot \mathrm{sigmaV} \cdot \mathrm{Jer} \tag{75}$$

# **6.76 Rule** g6p

Rule g6p is a rate rule for species g6p:

$$\frac{d}{dt}g6p = lambda \cdot (Rgk - pfk) \tag{76}$$

# **6.77 Rule** fbp

Rule fbp is a rate rule for species fbp:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{fbp} = \mathrm{lambda} \cdot \left(\frac{\mathrm{pfk}}{1} - 0.5 \cdot \mathrm{rgpdh}\right) \tag{77}$$

# **6.78 Rule** adp

Rule adp is a rate rule for species adp:

$$\frac{d}{dt}adp = \frac{atp - [adp] \cdot exp\left(fback \cdot \left(1 - \frac{[c]}{rI}\right)\right)}{taua \cdot 1}$$
(78)

# 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

Name V

**SBO:0000259** voltage

Initial amount -60 mol

Involved in rule V

One rule which determines this species' quantity.

# 7.2 Species n

Name n

Initial amount 0.025 mol

Involved in rule n

One rule which determines this species' quantity.

# 7.3 Species c

Name c

Initial amount 0.25 mol

Involved in rule c

One rule which determines this species' quantity.

# 7.4 Species cer

Name cer

Initial amount 185 mol

Involved in rule cer

One rule which determines this species' quantity.

# 7.5 Species g6p

Name g6p

Initial amount 200 mol

Involved in rule g6p

One rule which determines this species' quantity.

# 7.6 Species fbp

Name fbp

Initial amount 40 mol

Involved in rule fbp

One rule which determines this species' quantity.

# 7.7 Species adp

Name adp

Initial amount 780 mol

Involved in rule adp

One rule which determines this species' quantity.

# A Glossary of Systems Biology Ontology Terms

**SBO:000009 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 id the Farad

**SBO:0000259 voltage:** Difference of electrical potential between two points of an electrical network, expressed in volts

SML2ATEX was developed by Andreas Dräger<sup>a</sup>, Hannes Planatscher<sup>a</sup>, Dieudonné M Wouamba<sup>a</sup>, Adrian Schröder<sup>a</sup>, Michael Hucka<sup>b</sup>, Lukas Endler<sup>c</sup>, Martin Golebiewski<sup>d</sup> and Andreas Zell<sup>a</sup>. Please see http://www.ra.cs.uni-tuebingen.de/software/SBML2LaTeX for more information.

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

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

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

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