# **SBML Model Report**

# Model name: "Proctor2016 - Circadian rhythm of PTH and the dynamics of signaling molecules on bone remodeling"



August 25, 2016

## 1 General Overview

This is a document in SBML Level 2 Version 4 format. This model was created by the following two authors: Vijayalakshmi Chelliah<sup>1</sup> and Carole Proctor<sup>2</sup> at May 26<sup>th</sup> 2016 at 4:04 p.m. and last time modified at June nineth 2016 at 10:37 a.m. Table 1 provides an overview of the quantities of all components of this model.

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

Element	Quantity	Element	Quantity
compartment types	0	compartments	3
species types	0	species	37
events	4	constraints	0
reactions	76	function definitions	0
global parameters	72	unit definitions	1
rules	0	initial assignments	0

#### **Model Notes**

Proctor2016 - Circadian rhythm of PTH and thedynamics of signaling molecules on bone remodeling

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This model is described in the article: Simulated Interventions to Ameliorate Age-Related Bone Loss Indicate the Importance of Timing. Proctor CJ, Gartland A. Front Endocrinol (Lausanne) 2016; 7: 61

Abstract:

Bone remodeling is the continuous process of bone resorption by osteoclasts and bone formation by osteoblasts, in order to maintain homeostasis. The activity of osteoclasts and osteoblasts is regulated by a network of signaling pathways, including Wnt, parathyroid hormone (PTH), RANK ligand/osteoprotegrin, and TGF-?, in response to stimuli, such as mechanical loading. During aging there is a gradual loss of bone mass due to dysregulation of signaling pathways. This may be due to a decline in physical activity with age and/or changes in hormones and other signaling molecules. In particular, hormones, such as PTH, have a circadian rhythm, which may be disrupted in aging. Due to the complexity of the molecular and cellular networks involved in bone remodeling, several mathematical models have been proposed to aid understanding of the processes involved. However, to date, there are no models, which explicitly consider the effects of mechanical loading, the circadian rhythm of PTH, and the dynamics of signaling molecules on bone remodeling. Therefore, we have constructed a network model of the system using a modular approach, which will allow further modifications as required in future research. The model was used to simulate the effects of mechanical loading and also the effects of different interventions, such as continuous or intermittent administration of PTH. Our model predicts that the absence of regular mechanical loading and/or an impaired PTH circadian rhythm leads to a gradual decrease in bone mass over time, which can be restored by simulated interventions and that the effectiveness of some interventions may depend on their timing.

This model is hosted on BioModels Database and identified by: BIOMD0000000612.

To cite BioModels Database, please use: BioModels Database: An enhanced, curated and annotated resource for published quantitative kinetic models.

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## 2 Unit Definitions

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

#### 2.1 Unit substance

**Definition** item

#### 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 Compartments

This model contains three compartments.

Table 2: Properties of all compartments.

Id	Name	SBO	Spatial Dimensions	Size	Unit	Constant	Outside
bone			3	1	litre	$ \overline{\mathbf{Z}} $	
ecm			3	1	litre	$   \overline{\mathbf{Z}} $	
osteoblast			3	1	litre	$   \overline{\mathbf{Z}} $	

## 3.1 Compartment bone

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

## 3.2 Compartment ecm

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

## 3.3 Compartment osteoblast

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

# 4 Species

This model contains 37 species. The boundary condition of two of these species is set to true so that these species' amount cannot be changed by any reaction. 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
Bone		bone	item		
HSC		bone	item		
$0b_m$		bone	item		
Ob_p		bone	item		$\Box$
$Ob_p_Tgfb_A$		bone	item		$\Box$
Ob_pro		bone	item		$\Box$
Ocl_m		bone	item		
$0cl_p$		bone	item		
$Ocl_p_RANKL$		bone	item		$\Box$
$\mathtt{Ocy}_{-}\mathtt{A}$		bone	item		
Ocy_I		bone	item		$\Box$
$Ocy_I_PTH$		bone	item		
MSC		bone	item		$\Box$
LOAD		ecm	item		$\Box$
MCSF		ecm	item		
Ob_m_PTH		ecm	item		
$0b_p_TH$		ecm	item		
OPG		ecm	item		
OPG_RANKL		ecm	item		
PTH		ecm	item		
RANKL		ecm	item		

Id	Name	Compartment	Derived Unit	Constant	Boundary Condi- tion
Sost		ecm	item		
Tgfb_A		ecm	item		
${\sf Tgfb}_{\sf -I}$		ecm	item		
Wnt_A		ecm	item		
${ t Wnt_I}$		ecm	item		
Source		ecm	item		
Sink		ecm	item	$\overline{\mathbf{Z}}$	$   \overline{\checkmark} $
X		ecm	item		
newbone		ecm	item		
Bax		osteoblast	item		
Bax_Bcl2		osteoblast	item		
Bcl2		osteoblast	item		
CREB		osteoblast	item		
CREB_P		osteoblast	item		
CREB_Runx2		osteoblast	item		
Runx2		osteoblast	item		$\Box$

# **5 Parameters**

This model contains 72 global parameters.

Table 4: Properties of each parameter.

kactCreb         0.009         Z           kactOcy         0.004         Z           kactTgfb         2.10-7         Z           kactWnt         0.030         Z           kactWntPth         0.001         Z           kbinCebRunx2         0.010         Z           kbinCobpTH         0.020         Z           kbinObpTH         3.10-4         Z           kbinObpTgfb         2.10-4         Z           kbinOclpRANKL         0.001         Z           kbinOcpTH         0.008         Z           kdeathOb         2.4·10-4         Z           kdeathOcl         6.5·10-5         Z           kdeathOcl         10-8         Z           kdeathOcy         10-8         Z           kdegBCl2         0.003         Z           kdegBone         6.5·10-9         Z           kdegDGG         4·10-6         Z           kdegOPGRANKL         10-5         Z           kdegPAMKL         3·10-5         Z           kdegRanx2         10-4         Z           kdegRost         0.004         Z           kdegTfbTH         1.7·10-5         Z      <	Id	Name	SBO	Value	Unit	Constant
kactOcy         0.004         Image: square s	kactCreb			0.009		<b>✓</b>
kactTgfb         2·10 <sup>-7</sup> Z           kactWnt         0.030         Z           kactWntPth         0.001         Z           kbinBaxBc12         0.010         Z           kbinCrebRunx2         0.010         Z           kbinObpTH         0.020         Z           kbinObpTH         3·10 <sup>-4</sup> Z           kbinOclpRANKL         0.001         Z           kbinOclpRANKL         0.001         Z           kbinOclpRANKL         0.001         Z           kdeathOcl         6.5·10 <sup>-5</sup> Z           kdeathOcl         6.5·10 <sup>-5</sup> Z           kdeathOcl         10 <sup>-8</sup> Z           kdegBc12         0.003         Z           kdegBc2         0.003         Z           kdegDGG         4·10 <sup>-6</sup> Z           kdegDGG         4·10 <sup>-6</sup> Z           kdegPTH         0.002         Z           kdegRANKL         3·10 <sup>-5</sup> Z           kdegRankL         3·10 <sup>-5</sup> Z           kdegRank2         10 <sup>-4</sup> Z           kdegTgfb         5·10 <sup>-5</sup> Z           kdegTgfbTH         1.7·10 <sup>-5</sup>	kact0cy			0.004		
kactWnt         0.030         1           kactWntPth         0.001         1           kbinBaxBc12         0.010         2           kbinCrebRunx2         0.010         2           kbinObmPTH         0.020         2           kbinObpPTH         3 · 10 - 4         2           kbinOclpRANKL         0.001         2           kbinOclpRANKL         0.001         2           kdeathOcl         6.5 · 10 - 5         2           kdeathOcl         6.5 · 10 - 5         2           kdeathOcy         10 - 8         2           kdegBc12         0.003         2           kdegBc12         0.003         2           kdegDPG         4 · 10 - 6         2           kdegDPG         4 · 10 - 6         2           kdegPTH         0.002         2           kdegRanx2         10 - 4         2           kdegRanx2         10 - 4         2           kdegTgfb         5 · 10 - 5         2           kdegTgfb         5 · 10 - 5         2           kdefTMSC         5 · 5 · 10 - 5         2           kdiffObP         10 - 4         2           kdiffObP         0.050<	kact0cyPth	ı		0.080		
kactWntPth         0.001         7           kbinBaxBc12         0.010         7           kbinCrebRunx2         0.010         7           kbinObpTH         0.020         7           kbinObpTgfb         2·10-4         7           kbinOc1pRANKL         0.001         7           kbinOc2pTH         0.008         7           kdeathOb         2.4·10-4         7           kdeathOc1         6.5·10-5         7           kdeathOc2p         10-8         7           kdegBc12         0.003         7           kdegBc12         0.003         7           kdegDPG         4·10-6         7           kdegOPGRANKL         10-5         7           kdegPTH         0.002         7           kdegRanx2         10-4         7           kdegRanx2         10-4         7           kdegRost         0.003         7           kdegTgfb         5·10-5         7           kdegTgfbTH         1.7·10-5         7           kdegTgfbTH         1.7·10-5         7           kdegTgfbTH         1.7·10-5         7           kdegTgfbPTH         1.7·10-5         7	$\mathtt{kactTgfb}$			$2 \cdot 10^{-7}$		
kbinBaxBc12         0.010           kbinCrebRunx2         0.010           kbinObmPTH         0.020           kbinObpTgfb         2·10-4           kbinOclpRANKL         0.001           kbinOclpRANKL         0.008           kdeathOb         2.4·10-4           kdeathOclp         10-5           kdeathOclp         10-8           kdegBc12         0.003           kdegBone         6.5·10-9           kdegBone         6.5·10-9           kdegOPG         4·10-6           kdegPTH         0.002           kdegRANKL         3·10-5           kdegRunx2         10-4           kdegSost         0.004           kdegTgfb         5·10-5           kdegTgfbPTH         1.7·10-5           kdegTgfbPTH         1.7·10-5           kdiffMSC         6.5·10-4           kdiffObP         10-4           kdiffObP         10-4           kdiffObP         10-5           kdiffOtP         8·10-5           kdiffOtP         8·10-5           kformBone         3.07·10-6           kinactCreb         10-4	kactWnt			0.030		
kbinCrebRunx2         0.010           kbinObmPTH         0.020           kbinObpTfH         3 · 10 - 4           kbinObpTgfb         2 · 10 - 4           kbinOclpRANKL         0.001           kbinOcyPTH         0.008           kdeathOb         2 · 4 · 10 - 4           kdeathOcl         6 · 5 · 10 - 5           kdeathOcl         10 - 5           kdeathOcy         10 - 8           kdegBc12         0.003           kdegBone         6 · 5 · 10 - 9           kdegOPG         4 · 10 - 6           kdegOPGRANKL         10 - 5           kdegPTH         0.002           kdegRunx2         10 - 4           kdegRunx2         10 - 4           kdegSost         0.004           kdegTgfb         5 · 10 - 5           kdegTffBSC         5 · 5 · 10 - 5           kdiffBSC         6 · 5 · 10 - 4           kdiffObP         10 - 4           kdiffObP         10 - 4           kdiffOcIP         8 · 10 - 5           kinactCreb         10 - 4	kactWntPth	ı		0.001		
kbinObmPTH         0.020           kbinObpTfb         3 · 10^{-4}           kbinObpTgfb         2 · 10^{-4}           kbinOc1pRANKL         0.001           kbinOcyPTH         0.008           kdeathOb         2 · 4 · 10^{-4}           kdeathOc1         6 · 5 · 10^{-5}           kdeathOc2         10^{-5}           kdeathOcy         10^{-8}           kdegBc12         0.003           kdegBone         6 · 5 · 10^{-9}           kdegDPG         4 · 10^{-6}           kdegPTH         0.002           kdegRANKL         3 · 10^{-5}           kdegRunx2         10^{-4}           kdegRunx2PTH         0.003           kdegTgfb         5 · 10^{-5}           kdegTgfb         5 · 10^{-5}           kdegTgfb         5 · 10^{-5}           kdiffMSC         6 · 5 · 10^{-4}           kdiffObP         10^{-4}           kdiffOcP         0.050           kdiffOcIP         8 · 10^{-5}           kinactCreb         10^{-4}	kbinBaxBc]	L2		0.010		
kbinObpTff       3 · 10 - 4         kbinObpTgfb       2 · 10 - 4         kbinOclpRANKL       0.001         kbinOcyPTH       0.008         kdeathOb       2.4 · 10 - 4         kdeathOclp       10 - 5         kdeathOcy       10 - 8         kdegBc12       0.003         kdegBone       6.5 · 10 - 9         kdegOPG       4 · 10 - 6         kdegOPGRANKL       10 - 5         kdegPTH       0.002         kdegRankL       3 · 10 - 5         kdegRunx2       10 - 4         kdegSost       0.004         kdegTgfb       5 · 10 - 5         kdegTgfbFTH       1.7 · 10 - 5         kdefffMSC       5.5 · 10 - 5         kdiffMSC       6.5 · 10 - 4         kdiffObP       10 - 4         kdiffObproTgfb       0.050         kdiffOclP       8 · 10 - 5         kformBone       3.07 · 10 - 6         kinactCreb       10 - 4	kbinCrebRu	ınx2		0.010		
kbinObpTgfb       3 · 10^-4         kbinOclpRANKL       0.001         kbinOcyPTH       0.008         kdeathOb       2.4 · 10^-4         kdeathOcl       6.5 · 10^-5         kdeathOclp       10^-5         kdeathOcy       10^-8         kdegBc12       0.003         kdegBone       6.5 · 10^-9         kdegOPG       4 · 10^-6         kdegOPGRANKL       10^-5         kdegTH       0.002         kdegRankL       3 · 10^-5         kdegRunx2       10^-4         kdegSost       0.004         kdegTgfb       5 · 10^-5         kdegTgfbPTH       1.7 · 10^-5         kdegTgfbSC       5.5 · 10^-5         kdiffMSC       5.5 · 10^-5         kdiffObP       10^-4         kdiffObP       10^-4         kdiffOclP       8 · 10^-5         kdiffOclP       8 · 10^-5         kinactCreb       10^-4	kbin0bmPTH	I		0.020		_
kbinObpTgfb         2·10 <sup>-4</sup> kbinOclpRANKL         0.001           kbinOcyPTH         0.008           kdeathOb         2.4·10 <sup>-4</sup> kdeathOcl         6.5·10 <sup>-5</sup> kdeathOclp         10 <sup>-5</sup> kdeathOcy         10 <sup>-8</sup> kdegBc12         0.003           kdegBone         6.5·10 <sup>-9</sup> kdegOPG         4·10 <sup>-6</sup> kdegOPG         4·10 <sup>-6</sup> kdegPTH         0.002           kdegRankL         3·10 <sup>-5</sup> kdegRunx2         10 <sup>-4</sup> kdegSost         0.004           kdegTgfb         5·10 <sup>-5</sup> kdegTgfbPTH         1.7·10 <sup>-5</sup> kdiffHSC         5·5·10 <sup>-5</sup> kdiffMSC         6·5·10 <sup>-4</sup> kdiffObP         10 <sup>-4</sup> kdiffObP         10 <sup>-4</sup> kdiffOclP         8·10 <sup>-5</sup> kdiffOclP         8·10 <sup>-5</sup> kinactCreb         10 <sup>-4</sup>	kbinObpPTH	I		$3 \cdot 10^{-4}$		
kbinOcyPTH       0.008         kdeathOb       2.4 · 10^{-4}         kdeathOcl       6.5 · 10^{-5}         kdeathOclp       10^{-8}         kdeathOcy       10^{-8}         kdegBc12       0.003         kdegBone       6.5 · 10^{-9}         kdegOPG       4 · 10^{-6}         kdegOPGRANKL       10^{-5}         kdegPTH       0.002         kdegRankL       3 · 10^{-5}         kdegRunx2       10^{-4}         kdegRunx2PTH       0.003         kdegTgfb       5 · 10^{-5}         kdegTgfbPTH       1.7 · 10^{-5}         kdiffHSC       5.5 · 10^{-5}         kdiffMSC       6.5 · 10^{-4}         kdiffObP       10^{-4}         kdiffObPorOTgfb       0.050         kdiffOclP       8 · 10^{-5}         kformBone       3.07 · 10^{-6}         kinactCreb       10^{-4}	kbinObpTgf	b		$2 \cdot 10^{-4}$		
kbinOcyPTH       0.008         kdeathOb       2.4 · 10^{-4}         kdeathOcl       6.5 · 10^{-5}         kdeathOclp       10^{-8}         kdeathOcy       10^{-8}         kdegBc12       0.003         kdegBone       6.5 · 10^{-9}         kdegMCSF       10^{-4}         kdegOPG       4 · 10^{-6}         kdegOPGRANKL       10^{-5}         kdegPTH       0.002         kdegRANKL       3 · 10^{-5}         kdegRunx2       10^{-4}         kdegRunx2PTH       0.003         kdegTgfb       5 · 10^{-5}         kdegTgfbPTH       1.7 · 10^{-5}         kdiffHSC       5.5 · 10^{-5}         kdiffMSC       6.5 · 10^{-4}         kdiffObP       10^{-4}         kdiffOclP       8 · 10^{-5}         kdiffOclP       8 · 10^{-5}         kformBone       3.07 · 10^{-6}         kinactCreb       10^{-4}	kbinOclpRA	ANKL		0.001		
kdeath0b $2.4 \cdot 10^{-4}$ kdeath0cl $6.5 \cdot 10^{-5}$ kdeath0cy $10^{-8}$ kdegBc12 $0.003$ kdegBone $6.5 \cdot 10^{-9}$ kdegOPG $4 \cdot 10^{-6}$ kdegOPGRANKL $10^{-5}$ kdegPTH $0.002$ kdegRanx2 $10^{-4}$ kdegRunx2 $10^{-4}$ kdegSost $0.004$ kdegTgfb $5 \cdot 10^{-5}$ kdegTgfbPTH $1.7 \cdot 10^{-5}$ kdiffHSC $5.5 \cdot 10^{-5}$ kdiffMSC $6.5 \cdot 10^{-4}$ kdiffObP $10^{-4}$ kdiffOclP $8 \cdot 10^{-5}$ kdiffOclP $8 \cdot 10^{-5}$ kdiffOclP $0.050$ kdiffOclP $0.050$ kdiffOclP $0.050$ kdiffOclP $0.050$ kinactCreb $0.04$	kbin0cyPTH	I		0.008		_
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kdeathOcy $10^{-8}$ kdegBcl2 $0.003$ kdegBone $6.5 \cdot 10^{-9}$ kdegMCSF $10^{-4}$ kdegOPG $4 \cdot 10^{-6}$ kdegPTH $0.002$ kdegRANKL $3 \cdot 10^{-5}$ kdegRunx2 $10^{-4}$ kdegRunx2PTH $0.003$ kdegSost $0.004$ kdegTgfb $5 \cdot 10^{-5}$ kdegTgfbPTH $1.7 \cdot 10^{-5}$ kdiffHSC $5.5 \cdot 10^{-5}$ kdiffObP $10^{-4}$ kdiffObproTgfb $0.050$ kdiffOclP $8 \cdot 10^{-5}$ kformBone $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	kdeath0clp	)				_
kdegBole $6.5 \cdot 10^{-9}$ kdegMCSF $10^{-4}$ kdegOPG $4 \cdot 10^{-6}$ kdegPTH $0.002$ kdegRANKL $3 \cdot 10^{-5}$ kdegRunx2 $10^{-4}$ kdegSost $0.003$ kdegTgfb $5 \cdot 10^{-5}$ kdegTgfbPTH $1.7 \cdot 10^{-5}$ kdiffMSC $5.5 \cdot 10^{-5}$ kdiffObP $10^{-4}$ kdiffObPoroTgfb $0.050$ kdiffOclP $8 \cdot 10^{-5}$ kdiffOclP $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	kdeath0cy			$10^{-8}$		_
kdegMCSF $10^{-4}$ kdegOPG $4 \cdot 10^{-6}$ kdegOPGRANKL $10^{-5}$ kdegPTH $0.002$ kdegRANKL $3 \cdot 10^{-5}$ kdegRunx2 $10^{-4}$ kdegRunx2PTH $0.003$ kdegSost $0.004$ kdegTgfb $5 \cdot 10^{-5}$ kdiffHSC $5.5 \cdot 10^{-5}$ kdiffMSC $6.5 \cdot 10^{-4}$ kdiffObP $10^{-4}$ kdiffObP $10^{-4}$ kdiffOclP $8 \cdot 10^{-5}$ kformBone $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	kdegBcl2			0.003		_
kdegOPG $4 \cdot 10^{-6}$ kdegOPGRANKL $10^{-5}$ kdegPTH $0.002$ kdegRANKL $3 \cdot 10^{-5}$ kdegRunx2 $10^{-4}$ kdegSost $0.003$ kdegTgfb $5 \cdot 10^{-5}$ kdegTgfbPTH $1.7 \cdot 10^{-5}$ kdiffHSC $5.5 \cdot 10^{-5}$ kdiffMSC $6.5 \cdot 10^{-4}$ kdiffObP $10^{-4}$ kdiffOclP $8 \cdot 10^{-5}$ kdiffOclP $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	kdegBone			$6.5\cdot 10^{-9}$		
kdegOPGRANKL $10^{-6}$ kdegPTH $0.002$ kdegRANKL $3 \cdot 10^{-5}$ kdegRunx2 $10^{-4}$ kdegRunx2PTH $0.003$ kdegTgfb $5 \cdot 10^{-5}$ kdegTgfbPTH $1.7 \cdot 10^{-5}$ kdiffHSC $5.5 \cdot 10^{-5}$ kdiffMSC $6.5 \cdot 10^{-4}$ kdiffObP $10^{-4}$ kdiffObPoroTgfb $0.050$ kdiffOclP $8 \cdot 10^{-5}$ kformBone $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	kdegMCSF					
kdegOPGRANKL $10^{-5}$ kdegPTH $0.002$ kdegRANKL $3 \cdot 10^{-5}$ kdegRunx2 $10^{-4}$ kdegRunx2PTH $0.003$ kdegSost $0.004$ kdegTgfb $5 \cdot 10^{-5}$ kdegTgfbPTH $1.7 \cdot 10^{-5}$ kdiffHSC $5.5 \cdot 10^{-5}$ kdiffObP $10^{-4}$ kdiffObP $10^{-4}$ kdiffOclP $8 \cdot 10^{-5}$ kdiffOclP $8 \cdot 10^{-5}$ kformBone $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	kdegOPG			$4 \cdot 10^{-6}$		
kdegPTH $0.002$ kdegRANKL $3 \cdot 10^{-5}$ kdegRunx2 $10^{-4}$ kdegSost $0.004$ kdegTgfb $5 \cdot 10^{-5}$ kdegTgfbPTH $1.7 \cdot 10^{-5}$ kdiffHSC $5.5 \cdot 10^{-5}$ kdiffMSC $6.5 \cdot 10^{-4}$ kdiffObP $10^{-4}$ kdiffOclP $8 \cdot 10^{-5}$ kdiffOclP $8 \cdot 10^{-5}$ kformBone $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	kdegOPGRAN	IKL		$10^{-5}$		
kdegRANKL $3 \cdot 10^{-5}$ kdegRunx2 $10^{-4}$ kdegRunx2PTH $0.003$ kdegSost $0.004$ kdegTgfb $5 \cdot 10^{-5}$ kdegTgfbPTH $1.7 \cdot 10^{-5}$ kdiffHSC $5.5 \cdot 10^{-5}$ kdiffMSC $6.5 \cdot 10^{-4}$ kdiffObP $10^{-4}$ kdiffObproTgfb $0.050$ kdiffOclP $8 \cdot 10^{-5}$ kformBone $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	kdegPTH			0.002		
kdegRunx2 $10^{-4}$ kdegSost $0.004$ kdegTgfb $5 \cdot 10^{-5}$ kdegTgfbPTH $1.7 \cdot 10^{-5}$ kdiffHSC $5.5 \cdot 10^{-5}$ kdiffObP $10^{-4}$ kdiffObProTgfb $0.050$ kdiffOclP $8 \cdot 10^{-5}$ kformBone $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	${\tt kdegRANKL}$			$3 \cdot 10^{-5}$		_
kdegRunx2PTH $0.003$ kdegSost $0.004$ kdegTgfb $5 \cdot 10^{-5}$ kdegTgfbPTH $1.7 \cdot 10^{-5}$ kdiffHSC $5.5 \cdot 10^{-5}$ kdiffMSC $6.5 \cdot 10^{-4}$ kdiffObP $10^{-4}$ kdiffObproTgfb $0.050$ kdiffOclP $8 \cdot 10^{-5}$ kformBone $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	kdegRunx2			$10^{-4}$		_
kdegSost $0.004$ kdegTgfb $5 \cdot 10^{-5}$ kdegTgfbPTH $1.7 \cdot 10^{-5}$ kdiffHSC $5.5 \cdot 10^{-5}$ kdiffMSC $6.5 \cdot 10^{-4}$ kdiffObP $10^{-4}$ kdiffObproTgfb $0.050$ kdiffOclP $8 \cdot 10^{-5}$ kformBone $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	kdegRunx2F	PTH		0.003		
kdegTgfb $5 \cdot 10^{-5}$ kdegTgfbPTH $1.7 \cdot 10^{-5}$ kdiffHSC $5.5 \cdot 10^{-5}$ kdiffMSC $6.5 \cdot 10^{-4}$ kdiffObP $10^{-4}$ kdiffObproTgfb $0.050$ kdiffOclP $8 \cdot 10^{-5}$ kformBone $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	kdegSost			0.004		
kdiffHSC $5.5 \cdot 10^{-5}$ kdiffMSC $6.5 \cdot 10^{-4}$ kdiff0bP $10^{-4}$ kdiff0bproTgfb $0.050$ kdiff0clP $8 \cdot 10^{-5}$ kformBone $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$				$5\cdot 10^{-5}$		_
kdiffHSC $5.5 \cdot 10^{-5}$ kdiffMSC $6.5 \cdot 10^{-4}$ kdiffObP $10^{-4}$ kdiffObproTgfb $0.050$ kdiffOclP $8 \cdot 10^{-5}$ kformBone $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	kdegTgfbPT	ГН		$1.7\cdot10^{-5}$		
kdiffMSC $6.5 \cdot 10^{-4}$ kdiffObP $10^{-4}$ kdiffObproTgfb $0.050$ kdiffOclP $8 \cdot 10^{-5}$ kformBone $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	kdiffHSC			$5.5\cdot10^{-5}$		
kdiff0bP $10^{-4}$ kdiff0bproTgfb $0.050$ kdiff0clP $8 \cdot 10^{-5}$ kformBone $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	kdiffMSC			$6.5\cdot 10^{-4}$		
kdiffObproTgfb $0.050$ kdiffOclP $8 \cdot 10^{-5}$ kformBone $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	kdiff0bP			$10^{-4}$		
kdiffOclP $8 \cdot 10^{-5}$ kformBone $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	kdiff0bpro	Tgfb		0.050		
kformBone $3.07 \cdot 10^{-6}$ kinactCreb $10^{-4}$	kdiff0clP			$8 \cdot 10^{-5}$		
kinactCreb $10^{-4}$	kformBone		3			
	kinactCreb	)				
	kinact0cy			$2\cdot 10^{-5}$		

Id	Name	SBO	Value	Unit	Constant
kinactWnt			0.800		$\overline{Z}$
kinhibRANKL			0.001		$\overline{\mathbf{Z}}$
kmat0b			$2 \cdot 10^{-9}$		$\overline{\mathbf{Z}}$
kmat0bTgfb			$10^{-8}$		$\overline{\mathbf{Z}}$
krelBaxBcl2			0.500		$ \overline{\checkmark} $
krelCrebRunx2	2		0.010		$ \overline{\checkmark} $
krelObmPTH			0.005		
krelObpPTH			0.005		
krelObpTgfb			0.010		
krelOclpRANKI			0.001		
krelOcyPTH			0.005		
krelRANKL			0.001		
ksecMCSFbyMSC	C		$10^{-5}$		
ksecMCSFby0bm	n		$10^{-5}$		
ksecMCSFby0bp			$10^{-5}$		
ksecMCSFby0bp	oro		$10^{-5}$		
${\tt ksecOPGbyObm}$			$10^{-5}$		
ksec0PGby0bp			$2 \cdot 10^{-6}$		
ksec0PGby0bpF	PTH		$10^{-6}$		
ksecRANKLbyMS	SC		$10^{-6}$		$\square$
ksecRANKLbyOb	om		$10^{-7}$		$\square$
ksecRANKLbyOb	omPTH		$10^{-6}$		$\square$
ksecRANKLbyOb			$3 \cdot 10^{-6}$		$\square$
ksecRANKLbyOb			$2 \cdot 10^{-5}$		$\square$
ksecRANKLbyOb	-		$7 \cdot 10^{-6}$		
ksecRANKLbyOb			$4 \cdot 10^{-6}$		
ksecRANKLbyOc	•		$10^{-6}$		
ksecRANKLbyOc	cyI		$10^{-7}$		$\square$
ksecSost			$7.5 \cdot 10^{-4}$		
ksecTgfb			$5 \cdot 10^{-5}$		
ksynBcl2			0.005		$\square$
ksynPTH			0.020		$\mathbf{Z}_{\underline{\cdot}}$
ksynRunx2			0.005		$\mathbf{Z}_{\underline{\cdot}}$
ksynX			0.012		$\mathbf{Z}_{\underline{j}}$
kunload			$3.5 \cdot 10^{-4}$		$\square$

# 6 Events

This is an overview of four events. Each event is initiated whenever its trigger condition switches from false to true. A delay function postpones the effects of an event to a later time point. At the time of execution, an event can assign values to species, parameters or compartments if

these are not set to constant.

## **6.1 Event** AddLOAD1

**Trigger condition** 

(1)

**Assignment** 

$$[LOAD] = 1$$

(2)

## **6.2 Event** AddLOAD2

**Trigger condition** 

(3)

**Assignment** 

$$[LOAD] = 1$$

(4)

## **6.3 Event** AddPTH1

**Trigger condition** 

(5)

**Assignment** 

$$[PTH] = 150$$

(6)

## **6.4 Event** AddPTH2

**Trigger condition** 

(7)

**Assignments** 

$$[PTH] = 170$$

(8)

$$[X] = 0$$

(9)

# 7 Reactions

This model contains 76 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	Unloading		$LOAD \xrightarrow{LOAD} Sink$	
2	Osteocyte- _activation		$Ocy_I + LOAD \xrightarrow{Ocy_I, LOAD} Ocy_A + LOAD$	
3	Osteocyte- _activation_by- _PTH		$Ocy\_I\_PTH \xrightarrow{Ocy\_I\_PTH} Ocy\_A + PTH$	
4	Osteocyte- _inactivation		$Ocy\_A \xrightarrow{Ocy\_A} Ocy\_I$	
5	Ocy_I_bound_by- _PTH		$Ocy\_I + PTH \xrightarrow{Ocy\_I, PTH} Ocy\_I\_PTH$	
6	Ocy_I_PTH- _release		$Ocy\_I\_PTH \xrightarrow{Ocy\_I\_PTH} Ocy\_I + PTH$	
7	Ocy_apoptosis		$Ocy\_I \xrightarrow{Ocy\_I} Sink$	
8	Sost_secretion		$Ocy_{\bot} \xrightarrow{Ocy_{\bot} I} Ocy_{\bot} + Sost$	
9	Sost- _degradation		$Sost \xrightarrow{Sost} Sink$	
10	${\tt Wnt\_activation}$		$Wnt\_I \xrightarrow{Wnt\_I} Wnt\_A$	

N⁰	Id	Name	Reaction Equation	SBO
11	Wnt_activation- _by_PTH			
12	<pre>Inactivationof_Wnt_by_Sost</pre>		$Wnt\_A + Sost \xrightarrow{Wnt\_A, Sost} Wnt\_I + Sost$	
13	MSC- _differentiation- _to_Ob_pro	_	$MSC + Wnt\_A \xrightarrow{MSC, Wnt\_A} MSC + Wnt\_A + Ob\_pro$	
14	Osteoblast- _progenitor- _differentiation- _by_Tgfb	_	$Ob\_pro + Tgfb\_A \xrightarrow{Ob\_pro, Tgfb\_A} Ob\_p + Tgfb\_A$	
15	Ob_precursor- _differentiation		$Ob\_p \xrightarrow{Ob\_p} Ob\_m$	
16	Ob_p_binding- _by_PTH		$Ob_p + PTH \xrightarrow{Ob_p, PTH} Ob_p PTH$	
17	Ob_p_PTH- _release		$Ob\_p\_PTH \xrightarrow{Ob\_p\_PTH} Ob\_p + PTH$	
18	Ob_p_Tgfb- _binding		$Ob\_p + Tgfb\_A \xrightarrow{Ob\_p, Tgfb\_A} Ob\_p\_Tgfb\_A$	
19	Ob_p_Tgfb- _release		$Ob\_p\_Tgfb\_A \xrightarrow{Ob\_p\_Tgfb\_A} Ob\_p + Tgfb\_A$	
20	Ob_maturation- _to_Ocy		$Ob\_m \xrightarrow{Ob\_m} Ocy\_I$	

N⁰	Id Name	Reaction Equation SBO
21	Ob_maturation- _to_Ocy_by_Tgfb	$Ob\_m + Tgfb\_A \xrightarrow{Ob\_m, Tgfb\_A} Ocy\_I + Tgfb\_A$
22	Ob_m_bound_by- _PTH	$Ob\_m + PTH \xrightarrow{Ob\_m, PTH} Ob\_m\_PTH$
23	Ob_m_PTH- _release	$Ob\_m\_PTH \xrightarrow{Ob\_m\_PTH} Ob\_m + PTH$
24	Ob_m_apoptosis	$Ob_m + Bax \xrightarrow{Ob_m, Bax} Bax$
25	Ob_m_PTH- _apoptosis	$Ob\_m\_PTH + Bax \xrightarrow{Ob\_m\_PTH, Bax} Bax + PTH$
26	HSC- _differentiation- _to_Ocl_p	$HSC + MCSF \xrightarrow{HSC, MCSF} HSC + MCSF + Ocl_p$
27	Ocl_p_apoptosis	$Ocl_p \xrightarrow{Ocl_p} Sink$
28	Ocl_p_RANKL- _binding	$RANKL + Ocl_p \xrightarrow{Ocl_p, RANKL} Ocl_p\_RANKL$
29	Ocl_p_RANKL- _release	$Ocl\_p\_RANKL \xrightarrow{Ocl\_p\_RANKL} Ocl\_p + RANKL$
30	Osteoclastprecursordifferentiation	$Ocl\_p\_RANKL \xrightarrow{Ocl\_p\_RANKL} Ocl\_m$
31	RANKL- _inhibition	$OPG + RANKL \xrightarrow{OPG, RANKL} OPG\_RANKL$
32	OPG_RANKL- _dissociation	$OPG\_RANKL \xrightarrow{OPG\_RANKL} OPG + RANKL$

N⁰	Id	Name	Reaction Equation	SBO
33	OPG_RANKL- _degradation		OPG_RANKL OPG_RANKL Sink	
34	Osteoclast- _apoptosis		$Ocl_m \xrightarrow{Ocl_m} Sink$	
35	RANKL- _degradation		$RANKL \xrightarrow{RANKL} Sink$	
36	${\tt OPG\_degradation}$		$OPG \xrightarrow{OPG} Sink$	
37	MCSF_secretion- _by_MSC		$MSC \xrightarrow{MSC} MSC + MCSF$	
38	MCSF_secretion- _by_Ob_pro		$Ob\_pro \xrightarrow{Ob\_pro} Ob\_pro + MCSF$	
39	MCSF_secretion- _by_Ob_p		$Ob_{-}p \xrightarrow{Ob_{-}p} Ob_{-}p + MCSF$	
40	MCSF_secretion- _by_Ob_p_PTH		$Ob\_p\_PTH \xrightarrow{Ob\_p\_PTH} Ob\_p\_PTH + MCSF$	
41	MCSF_secretion- _by_Ob_m		$Ob\_m \xrightarrow{Ob\_m} Ob\_m + MCSF$	
42	MCSF_secretion- _by_Ob_m_PTH		$Ob\_m\_PTH \xrightarrow{Ob\_m\_PTH} Ob\_m\_PTH + MCSF$	
43	MCSF- _degradation		$MCSF \xrightarrow{MCSF} Sink$	
44	OPG_secretion- _by_Ob_p		$Ob_{-}p \xrightarrow{Ob_{-}p} Ob_{-}p + OPG$	

N₀	Id	Name	Reaction Equation	SBO
45	OPG_secretion- _by_Ob_p_PTH		$Ob\_p\_PTH \xrightarrow{Ob\_p\_PTH} Ob\_p\_PTH + OPG$	
46	OPG_secretion- _by_Ob_m		$Ob\_m \xrightarrow{Ob\_m} Ob\_m + OPG$	
47	RANKL- _secretion- _by_Ocy_A		$Ocy\_A \xrightarrow{Ocy\_A} Ocy\_A + RANKL$	
48	RANKL- _secretion- _by_Ocy_I		$Ocy\_I \xrightarrow{Ocy\_I} Ocy\_I + RANKL$	
49	RANKL- _secretion- _by_MSCs		$MSC \xrightarrow{MSC} MSC + RANKL$	
50	RANKL- _secretion- _by_Ob_p		$Ob\_p \xrightarrow{Ob\_p} Ob\_p + RANKL$	
51	RANKL- _secretion- _by_Ob_p_Tgfb_A		$Ob\_p\_Tgfb\_A \xrightarrow{Ob\_p\_Tgfb\_A} Ob\_p\_Tgfb\_A + RANKL$	-
52	RANKL- _secretion- _by_Ob_p_PTH		$Ob\_p\_PTH \xrightarrow{Ob\_p\_PTH} Ob\_p\_PTH + RANKL$	
53	RANKL- _secretion- _by_Ob_pro		$Ob\_pro \xrightarrow{Ob\_pro} Ob\_pro + RANKL$	

Produced by SBMIZIFEX	N₀	Id	Name	Reaction Equation	SBO
	54	RANKL- _secretion- _by_Ob_m		$Ob\_m \xrightarrow{Ob\_m} Ob\_m + RANKL$	
	55	RANKLsecretionby_Ob_m_PTHenhanced		$Ob\_m\_PTH \xrightarrow{Ob\_m\_PTH} Ob\_m\_PTH + RANKL$	
	56	Tgfb_secretion- _by_Obm		$Ob\_m \xrightarrow{Ob\_m} Ob\_m + Tgfb\_I$	
	57	${\tt Tgfb\_activation}$		$Tgfb\_I + Ocl\_m \xrightarrow{Tgfb\_I, Ocl\_m} Tgfb\_A + Ocl\_m$	
	58	Tgfb- _degradation		$Tgfb\_A \xrightarrow{Tgfb\_A} Sink$	
	59	Tgfb- _degradation- _by_PTH		$Tgfb\_A + Ob\_m\_PTH \xrightarrow{Tgfb\_A, Ob\_m\_PTH} Ob\_m\_PTH$	H
	60	$PTH\_production$		Source $\xrightarrow{\text{Source}}$ PTH	
	61	$PTH\_degradation$		$PTH \xrightarrow{PTH} Sink$	
	62	CREB_activation		$\begin{array}{c} \text{Ob\_m\_PTH} + \text{CREB} \xrightarrow{\text{CREB, Ob\_m\_PTH}} \text{Ob\_m\_PTH} - \\ \text{CREB\_P} \end{array}$	H
	63	CREB- _inactivation		$CREB\_P \xrightarrow{CREB\_P} CREB$	
	64	CREB_Runx2- _binding		$CREB\_P + Runx2 \xrightarrow{CREB\_P, Runx2} CREB\_Runx2$	

N⁰	Id	Name	Reaction Equation	SBO
65	CREB_Runx2- _release		CREB_Runx2 CREB_Runx2 CREB_P + Runx2	
66	Bcl2_synthesis		$CREB\_Runx2 \xrightarrow{CREB\_Runx2} CREB\_Runx2 + Bcl2$	
67	Bcl2- _degradation		$Bcl2 \xrightarrow{Bcl2} Sink$	
68	Bax_Bcl2- _binding		$Bax + Bcl2 \xrightarrow{Bax, Bcl2} Bax\_Bcl2$	
69	Bax_Bcl2- _release		$Bax\_Bcl2 \xrightarrow{Bax\_Bcl2} Bax + Bcl2$	
70	$Runx2\_synthesis$		Source $\xrightarrow{\text{Source}} \text{Runx}2$	
71	Runx2- _degradation- _via_PTH		$Ob\_m\_PTH + Runx2 \xrightarrow{Runx2, Ob\_m\_PTH} Ob\_m\_PTH$	
72	Runx2- _degradation		$Runx2 \xrightarrow{Runx2} Sink$	
73	Bone_formation		$Ob_m \xrightarrow{Ob_m} Ob_m + Bone + newbone$	
74	Bone_formation- _Obm_PTH		Ob_m_PTH Ob_m_PTH + Bone + newbone	
75	Bone- _degradation		$Ocl_m + Bone \xrightarrow{Ocl_m, Bone} Ocl_m$	
76	Xsynthesis		Source $\xrightarrow{\text{Source}} X$	

## 7.1 Reaction Unloading

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

## **Reaction equation**

$$LOAD \xrightarrow{LOAD} Sink \tag{10}$$

#### Reactant

Table 6: Properties of each reactant.

Id	Name	SBO
LOAD		

#### **Modifier**

Table 7: Properties of each modifier.

Id	Name	SBO
LOAD		

#### **Product**

Table 8: Properties of each product.

Id	Name	SBO
Sink		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_1 = \text{kunload} \cdot \text{LOAD}$$
 (11)

## 7.2 Reaction Osteocyte\_activation

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

## **Reaction equation**

$$Ocy\_I + LOAD \xrightarrow{Ocy\_I, LOAD} Ocy\_A + LOAD$$
 (12)

## **Reactants**

Table 9: Properties of each reactant.

Id	Name	SBO
Ocy_I		
LOAD		

#### **Modifiers**

Table 10: Properties of each modifier.

Id	Name	SBO
Ocy_I LOAD		

#### **Products**

Table 11: Properties of each product.

Name	SBO
	Name

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_2 = \text{kactOcy} \cdot \text{Ocy} \cdot \text{I} \cdot \text{LOAD}$$
 (13)

## 7.3 Reaction Osteocyte\_activation\_by\_PTH

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

## **Reaction equation**

$$Ocy\_I\_PTH \xrightarrow{Ocy\_I\_PTH} Ocy\_A + PTH$$
 (14)

#### Reactant

Table 12: Properties of each reactant.

Id	Name	SBO
Ocy_I_PTH		

## **Modifier**

Table 13: Properties of each modifier.

Id	Name	SBO
Ocy_I_PTH		

## **Products**

Table 14: Properties of each product.

Id	Name	SBO
Ocy_A PTH		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_3 = \text{kactOcyPth} \cdot \text{Ocy\_I\_PTH}$$
 (15)

## **7.4 Reaction** Osteocyte\_inactivation

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

## **Reaction equation**

$$Ocy\_A \xrightarrow{Ocy\_A} Ocy\_I \tag{16}$$

#### Reactant

Table 15: Properties of each reactant.

Id	Name	SBO
Ocy_A		

## **Modifier**

Table 16: Properties of each modifier.

Id	Name	SBO
Ocy_A		

## **Product**

Table 17: Properties of each product.

Id	Name	SBO
Ocy_I		

## **Kinetic Law**

Derived unit contains undeclared units

$$v_4 = \text{kinactOcy} \cdot \text{Ocy}_A$$
 (17)

## 7.5 Reaction Ocy\_I\_bound\_by\_PTH

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

## **Reaction equation**

$$Ocy_{-}I + PTH \xrightarrow{Ocy_{-}I, PTH} Ocy_{-}I_{-}PTH$$
(18)

## **Reactants**

Table 18: Properties of each reactant.

Id	Name	SBO
Ocy_I PTH		

#### **Modifiers**

Table 19: Properties of each modifier.

Id	Name	SBO
Ocy_I		
PTH		

#### **Product**

Table 20: Properties of each product.

Id	Name	SBO
Ocy_I_PTH		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_5 = \text{kbinOcyPTH} \cdot \text{Ocy} \cdot \frac{\text{PTH}^2}{100^2 + \text{PTH}^2}$$
 (19)

## 7.6 Reaction Ocy\_I\_PTH\_release

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

## **Reaction equation**

$$Ocy\_I\_PTH \xrightarrow{Ocy\_I\_PTH} Ocy\_I + PTH$$
 (20)

#### Reactant

Table 21: Properties of each reactant.

Id	Name	SBO
Ocy_I_PTH		

#### **Modifier**

Table 22: Properties of each modifier.

Id	Name	SBO
Ocy_I_PTH		

## **Products**

Table 23: Properties of each product.

Id	Name	SBO
Ocy_I PTH		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_6 = \text{krelOcyPTH} \cdot \text{Ocy\_I\_PTH}$$
 (21)

## 7.7 Reaction Ocy\_apoptosis

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

## **Reaction equation**

$$Ocy_{-}I \xrightarrow{Ocy_{-}I} Sink$$
 (22)

#### Reactant

Table 24: Properties of each reactant.

Id	Name	SBO
Ocy_I		

## **Modifier**

Table 25: Properties of each modifier.

Id	Name	SBO
Ocy_I		

#### **Product**

Table 26: Properties of each product.

Id	Name	SBO
Sink		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_7 = \text{kdeathOcy} \cdot \text{Ocy} \cdot \text{I}$$
 (23)

## 7.8 Reaction Sost\_secretion

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

## **Reaction equation**

$$Ocy \bot \xrightarrow{Ocy \bot} Ocy \bot + Sost$$
 (24)

## Reactant

Table 27: Properties of each reactant.

Id	Name	SBO
Ocy_I		

#### **Modifier**

Table 28: Properties of each modifier.

Id	Name	SBO
Ocy_I		

## **Products**

Table 29: Properties of each product.

Id	Name	SBO
Ocy_I		

Id	Name	SBO
Sost		

**Derived unit** contains undeclared units

$$v_8 = \text{ksecSost} \cdot \text{Ocy\_I}$$
 (25)

## 7.9 Reaction Sost\_degradation

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

## **Reaction equation**

$$Sost \xrightarrow{Sost} Sink \tag{26}$$

#### Reactant

Table 30: Properties of each reactant.

Id	Name	SBO
Sost		

## **Modifier**

Table 31: Properties of each modifier.

Id	Name	SBO
Sost		

## **Product**

Table 32: Properties of each product.

Id	Name	SBO
Sink		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_9 = \text{kdegSost} \cdot \text{Sost}$$
 (27)

#### 7.10 Reaction Wnt\_activation

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

## **Reaction equation**

$$Wnt\_I \xrightarrow{Wnt\_I} Wnt\_A \tag{28}$$

## Reactant

Table 33: Properties of each reactant.

Id	Name	SBO
Wnt_I		

#### **Modifier**

Table 34: Properties of each modifier.

Id	Name	SBO
$ootnotesize Wnt_I$		

#### **Product**

Table 35: Properties of each product.

Id	Name	SBO
Wnt_A		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{10} = \text{kactWnt} \cdot \text{Wnt}_{-}\text{I}$$
 (29)

## 7.11 Reaction Wnt\_activation\_by\_PTH

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

## **Reaction equation**

$$Wnt\_I + Ob\_m\_PTH \xrightarrow{Wnt\_I, Ob\_m\_PTH} Wnt\_A + Ob\_m\_PTH$$
(30)

## **Reactants**

Table 36: Properties of each reactant.

Id	Name	SBO
${ t Wnt_{-}I}$		
$\tt Ob\_m\_PTH$		

#### **Modifiers**

Table 37: Properties of each modifier.

Id	Name	SBO
Wnt_I		
$Ob_m_PTH$		

#### **Products**

Table 38: Properties of each product.

Id	Name	SBO
Wnt_A		
Ob_m_PTH		

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{11} = \text{kactWntPth} \cdot \text{Wnt\_I} \cdot \text{Ob\_m\_PTH}$$
 (31)

## **7.12 Reaction** Inactivation\_of\_Wnt\_by\_Sost

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

## **Reaction equation**

$$Wnt\_A + Sost \xrightarrow{Wnt\_A, Sost} Wnt\_I + Sost$$
 (32)

## **Reactants**

Table 39: Properties of each reactant.

Id	Name	SBO
Wnt_A		
Sost		

#### **Modifiers**

Table 40: Properties of each modifier.

Id	Name	SBO
Wnt_A		
Sost		

#### **Products**

Table 41: Properties of each product.

Id	Name	SBO
$\mathtt{Wnt}_{-}\mathbf{I}$		
Sost		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{12} = \frac{\text{kinactWnt} \cdot \text{Wnt\_A} \cdot \text{Sost}^2}{50^2 + \text{Sost}^2}$$
(33)

## 7.13 Reaction MSC\_differentiation\_to\_Ob\_pro

This is an irreversible reaction of two reactants forming three products influenced by two modifiers.

## **Reaction equation**

$$MSC + Wnt\_A \xrightarrow{MSC, Wnt\_A} MSC + Wnt\_A + Ob\_pro$$
 (34)

#### **Reactants**

Table 42: Properties of each reactant.

Id	Name	SBO
MSC		
${\tt Wnt\_A}$		

## **Modifiers**

Table 43: Properties of each modifier.

Id	Name	SBO
MSC		
${\tt Wnt\_A}$		

## **Products**

Table 44: Properties of each product.

Id	Name	SBO
MSC		
$\mathtt{Wnt}_{-}\mathtt{A}$		
Ob_pro		

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{13} = \frac{\text{kdiffMSC} \cdot \text{MSC} \cdot \text{Wnt\_A}^2}{50^2 + \text{Wnt\_A}^2}$$
(35)

## **7.14 Reaction** Osteoblast\_progenitor\_differentiation\_by\_Tgfb

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

## **Reaction equation**

$$Ob\_pro + Tgfb\_A \xrightarrow{Ob\_pro, Tgfb\_A} Ob\_p + Tgfb\_A$$
 (36)

#### **Reactants**

Table 45: Properties of each reactant.

Id	Name	SBO
Ob_pro		
Tgfb_A		

## **Modifiers**

Table 46: Properties of each modifier.

Id	Name	SBO
Ob_pro		
${\tt Tgfb\_A}$		

#### **Products**

Table 47: Properties of each product.

Id	Name	SBO
0b_p		
${\tt Tgfb\_A}$		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{14} = \frac{\text{kdiffObproTgfb} \cdot \text{Ob\_pro} \cdot \text{Tgfb\_A}^2}{50^2 + \text{Tgfb\_A}^2}$$
(37)

# **7.15 Reaction** Ob\_precursor\_differentiation

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

## **Reaction equation**

$$Ob\_p \xrightarrow{Ob\_p} Ob\_m$$
 (38)

#### Reactant

Table 48: Properties of each reactant.

Id	Name	SBO
Ob_p		

## **Modifier**

Table 49: Properties of each modifier.

Id	Name	SBO
0b_p		

## **Product**

Table 50: Properties of each product.

Id	Name	SBO
Ob_m		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{15} = \text{kdiffObP} \cdot \text{Ob\_p} \tag{39}$$

# 7.16 Reaction Ob\_p\_binding\_by\_PTH

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

## **Reaction equation**

$$Ob\_p + PTH \xrightarrow{Ob\_p, PTH} Ob\_p\_PTH$$
 (40)

## **Reactants**

Table 51: Properties of each reactant.

Id	Name	SBO
Ob_p PTH		

#### **Modifiers**

Table 52: Properties of each modifier.

Id	Name	SBO
Ob_p PTH		

#### **Product**

Table 53: Properties of each product.

Id	Name	SBO
Ob_p_PTH		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{16} = \text{kbinObpPTH} \cdot \text{Ob\_p} \cdot \frac{\text{PTH}^2}{100^2 + \text{PTH}^2}$$
 (41)

# 7.17 Reaction Ob\_p\_PTH\_release

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

## **Reaction equation**

$$Ob\_p\_PTH \xrightarrow{Ob\_p\_PTH} Ob\_p + PTH$$
 (42)

#### Reactant

Table 54: Properties of each reactant.

Id	Name	SBO
Ob_p_PTH		

#### **Modifier**

Table 55: Properties of each modifier.

Id	Name	SBO
Ob_p_PTH		

## **Products**

Table 56: Properties of each product.

Id	Name	SBO
Ob_p PTH		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{17} = \text{krelObpPTH} \cdot \text{Ob\_p\_PTH}$$
 (43)

## 7.18 Reaction Ob\_p\_Tgfb\_binding

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

## **Reaction equation**

$$Ob\_p + Tgfb\_A \xrightarrow{Ob\_p, Tgfb\_A} Ob\_p\_Tgfb\_A$$
 (44)

#### **Reactants**

Table 57: Properties of each reactant.

Id	Name	SBO
0b_p		
${\tt Tgfb\_A}$		

#### **Modifiers**

Table 58: Properties of each modifier.

Id	Name	SBO
Ob_p		

Id	Name	SBO
Tgfb_A		

## **Product**

Table 59: Properties of each product.

Id	Name	SBO
Ob_p_Tgfb_A		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{18} = kbinObpTgfb \cdot Ob_p \cdot Tgfb_A$$
 (45)

## 7.19 Reaction Ob\_p\_Tgfb\_release

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

## **Reaction equation**

$$Ob\_p\_Tgfb\_A \xrightarrow{Ob\_p\_Tgfb\_A} Ob\_p + Tgfb\_A$$
 (46)

#### Reactant

Table 60: Properties of each reactant.

Id	Name	SBO
Ob_p_Tgfb_A		

#### **Modifier**

Table 61: Properties of each modifier.

Id	Name	SBO
Ob_p_Tgfb_A		

## **Products**

Table 62: Properties of each product.

Id	Name	SBO
Ob_p		
Tgfb_A		

**Derived unit** contains undeclared units

$$v_{19} = krelObpTgfb \cdot Ob_p\_Tgfb\_A$$
 (47)

## **7.20 Reaction** Ob\_maturation\_to\_Ocy

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

## **Reaction equation**

$$Ob\_m \xrightarrow{Ob\_m} Ocy\_I$$
 (48)

## Reactant

Table 63: Properties of each reactant.

Id	Name	SBO
Ob_m		

#### **Modifier**

Table 64: Properties of each modifier.

Id	Name	SBO
Ob_m		

#### **Product**

Table 65: Properties of each product.

Id	Name	SBO
Ocy_I		

**Derived unit** contains undeclared units

$$v_{20} = \text{kmatOb} \cdot \text{Ob\_m} \tag{49}$$

## 7.21 Reaction Ob\_maturation\_to\_Ocy\_by\_Tgfb

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

## **Reaction equation**

$$Ob\_m + Tgfb\_A \xrightarrow{Ob\_m, Tgfb\_A} Ocy\_I + Tgfb\_A$$
 (50)

#### **Reactants**

Table 66: Properties of each reactant.

Id	Name	SBO
Ob_m		
$\mathtt{Tgfb}_{-}\mathtt{A}$		

#### **Modifiers**

Table 67: Properties of each modifier.

Id	Name	SBO
Ob_m		
${\tt Tgfb\_A}$		

#### **Products**

Table 68: Properties of each product.

Id	Name	SBO
Ocy_I		
${\tt Tgfb\_A}$		

**Derived unit** contains undeclared units

$$v_{21} = \frac{\text{kmatObTgfb} \cdot \text{Ob\_m} \cdot \text{Tgfb\_A}^2}{50^2 + \text{Tgfb\_A}^2}$$
 (51)

## 7.22 Reaction Ob\_m\_bound\_by\_PTH

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

## **Reaction equation**

$$Ob\_m + PTH \xrightarrow{Ob\_m, PTH} Ob\_m\_PTH$$
 (52)

#### **Reactants**

Table 69: Properties of each reactant.

Id	Name	SBO
Ob_m		
PTH		

## **Modifiers**

Table 70: Properties of each modifier.

Id	Name	SBO
Ob_m		
PTH		

## **Product**

Table 71: Properties of each product.

Id	Name	SBO
Ob_m_PTH		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{22} = \text{kbinObmPTH} \cdot \text{Ob\_m} \cdot \frac{\text{PTH}^2}{100^2 + \text{PTH}^2}$$
 (53)

## 7.23 Reaction Ob\_m\_PTH\_release

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

## **Reaction equation**

$$Ob\_m\_PTH \xrightarrow{Ob\_m\_PTH} Ob\_m + PTH$$
 (54)

## Reactant

Table 72: Properties of each reactant.

Id	Name	SBO
Ob_m_PTH		

#### **Modifier**

Table 73: Properties of each modifier.

Id	Name	SBO
Ob_m_PTH		

## **Products**

Table 74: Properties of each product.

Id	Name	SBO
Ob_m		
PTH		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{23} = \text{krelObmPTH} \cdot \text{Ob\_m\_PTH}$$
 (55)

## **7.24 Reaction** Ob\_m\_apoptosis

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

# **Reaction equation**

$$Ob_{-}m + Bax \xrightarrow{Ob_{-}m, Bax} Bax$$
 (56)

## **Reactants**

Table 75: Properties of each reactant.

Id	Name	SBO
Ob_m		
Bax		

#### **Modifiers**

Table 76: Properties of each modifier.

Id	Name	SBO
Ob_m		
Bax		

#### **Product**

Table 77: Properties of each product.

Id	Name	SBO
Bax		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{24} = \frac{\text{kdeathOb} \cdot \text{Ob} \cdot \text{m} \cdot \text{Bax}^2}{50^2 + \text{Bax}^2}$$
 (57)

# **7.25 Reaction** Ob\_m\_PTH\_apoptosis

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

## **Reaction equation**

$$Ob\_m\_PTH + Bax \xrightarrow{Ob\_m\_PTH, Bax} Bax + PTH$$
 (58)

## **Reactants**

Table 78: Properties of each reactant.

Id	Name	SBO
Ob_m_PTH		
Bax		

#### **Modifiers**

Table 79: Properties of each modifier.

Id	Name	SBO
Ob_m_PTH		
Bax		

#### **Products**

Table 80: Properties of each product.

Id	Name	SBO
Bax		
PTH		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{25} = \frac{\text{kdeathOb} \cdot \text{Ob\_m\_PTH} \cdot \text{Bax}^2}{50^2 + \text{Bax}^2}$$
 (59)

# **7.26 Reaction** HSC\_differentiation\_to\_Ocl\_p

This is an irreversible reaction of two reactants forming three products influenced by two modifiers.

## **Reaction equation**

$$HSC + MCSF \xrightarrow{HSC, MCSF} HSC + MCSF + Ocl_p$$
 (60)

#### **Reactants**

Table 81: Properties of each reactant.

Id	Name	SBO
HSC		
MCSF		

## **Modifiers**

Table 82: Properties of each modifier.

Id	Name	SBO
HSC		
MCSF		

## **Products**

Table 83: Properties of each product.

Id	Name	SBO
HSC		
MCSF		
Ocl_p		

### **Kinetic Law**

Derived unit contains undeclared units

$$v_{26} = \frac{\text{kdiffHSC} \cdot \text{HSC} \cdot \text{MCSF}^2}{50^2 + \text{MCSF}^2}$$
 (61)

# **7.27 Reaction** Ocl\_p\_apoptosis

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

# **Reaction equation**

$$Ocl\_p \xrightarrow{Ocl\_p} Sink$$
 (62)

## Reactant

Table 84: Properties of each reactant.

Id	Name	SBO
Ocl_p		

## **Modifier**

Table 85: Properties of each modifier.

Id	Name	SBO
Ocl_p		

## **Product**

Table 86: Properties of each product.

Id	Name	SBO
Sink		

## **Kinetic Law**

Derived unit contains undeclared units

$$v_{27} = \text{kdeathOclp} \cdot \text{Ocl}_{-p}$$
 (63)

# 7.28 Reaction Ocl\_p\_RANKL\_binding

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

# **Reaction equation**

$$RANKL + Ocl_{-p} \xrightarrow{Ocl_{-p}, RANKL} Ocl_{-p} \xrightarrow{RANKL}$$
 (64)

# **Reactants**

Table 87: Properties of each reactant.

Id	Name	SBO
RANKL		
$0cl_p$		

## **Modifiers**

Table 88: Properties of each modifier.

Id	Name	SBO
Ocl_p RANKL		

#### **Product**

Table 89: Properties of each product.

Id	Name	SBO
Ocl_p_RANKL		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{28} = \text{kbinOclpRANKL} \cdot \text{Ocl\_p} \cdot \text{RANKL}$$
 (65)

# 7.29 Reaction Ocl\_p\_RANKL\_release

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

# **Reaction equation**

$$Ocl_p\_RANKL \xrightarrow{Ocl\_p\_RANKL} Ocl\_p + RANKL$$
 (66)

# Reactant

Table 90: Properties of each reactant.

Id	Name	SBO
Ocl_p_RANKL		

#### **Modifier**

Table 91: Properties of each modifier.

Id	Name	SBO
Ocl_p_RANKL		

## **Products**

Table 92: Properties of each product.

Id	Name	SBO
Ocl_p RANKL		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{29} = \text{krelOclpRANKL} \cdot \text{Ocl\_p\_RANKL}$$
 (67)

# **7.30 Reaction** Osteoclast\_precursor\_differentiation

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

# **Reaction equation**

$$Ocl\_p\_RANKL \xrightarrow{Ocl\_p\_RANKL} Ocl\_m$$
 (68)

## Reactant

Table 93: Properties of each reactant.

Id	Name	SBO
Ocl_p_RANKL		

## **Modifier**

Table 94: Properties of each modifier.

Id	Name	SBO
Ocl_p_RANKL		

# **Product**

Table 95: Properties of each product.

Id	Name	SBO
Ocl_m		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{30} = \text{kdiffOclP} \cdot \text{Ocl\_p\_RANKL}$$
 (69)

## 7.31 Reaction RANKL\_inhibition

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

# **Reaction equation**

$$OPG + RANKL \xrightarrow{OPG, RANKL} OPG\_RANKL$$
 (70)

# **Reactants**

Table 96: Properties of each reactant.

Id	Name	SBO
OPG		
RANKL		

#### **Modifiers**

Table 97: Properties of each modifier.

Id	Name	SBO
OPG		
RANKL		

### **Product**

Table 98: Properties of each product.

Id	Name	SBO
OPG_RANKL		

**Derived unit** contains undeclared units

$$v_{31} = kinhibRANKL \cdot OPG \cdot RANKL \tag{71}$$

# 7.32 Reaction OPG\_RANKL\_dissociation

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

# **Reaction equation**

$$OPG\_RANKL \xrightarrow{OPG\_RANKL} OPG + RANKL$$
 (72)

#### Reactant

Table 99: Properties of each reactant.

Id	Name	SBO
OPG_RANKL		

## **Modifier**

Table 100: Properties of each modifier.

Id	Name	SBO
OPG_RANKL		

#### **Products**

Table 101: Properties of each product.

Id	Name	SBO
OPG		
RANKL		

**Derived unit** contains undeclared units

$$v_{32} = \text{krelRANKL} \cdot \text{OPG\_RANKL} \tag{73}$$

# 7.33 Reaction OPG\_RANKL\_degradation

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

# **Reaction equation**

$$OPG\_RANKL \xrightarrow{OPG\_RANKL} Sink$$
 (74)

#### Reactant

Table 102: Properties of each reactant.

Id	Name	SBO
OPG_RANKL	·	

### **Modifier**

Table 103: Properties of each modifier.

Id	Name	SBO
OPG_RANKL		

#### **Product**

Table 104: Properties of each product.

Id	Name	SBO
Sink		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{33} = \text{kdegOPGRANKL} \cdot \text{OPG\_RANKL}$$
 (75)

# **7.34 Reaction** Osteoclast\_apoptosis

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

# **Reaction equation**

$$Ocl_m \xrightarrow{Ocl_m} Sink \tag{76}$$

#### Reactant

Table 105: Properties of each reactant.

Id	Name	SBO
Ocl_m		

#### **Modifier**

Table 106: Properties of each modifier.

Id	Name	SBO
Ocl_m		

#### **Product**

Table 107: Properties of each product.

Id	Name	SBO
Sink		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{34} = \text{kdeathOcl} \cdot \text{Ocl}_{\text{m}}$$
 (77)

# 7.35 Reaction RANKL\_degradation

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

# **Reaction equation**

$$RANKL \xrightarrow{RANKL} Sink$$
 (78)

## Reactant

Table 108: Properties of each reactant.

Id	Name	SBO
RANKL		

## **Modifier**

Table 109: Properties of each modifier.

Id	Name	SBO
RANKL		

## **Product**

Table 110: Properties of each product.

Id	Name	SBO
Sink		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{35} = kdegRANKL \cdot RANKL \tag{79}$$

# **7.36 Reaction** OPG\_degradation

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

# **Reaction equation**

$$OPG \xrightarrow{OPG} Sink \tag{80}$$

### Reactant

Table 111: Properties of each reactant.

Id	Name	SBO
OPG		

### **Modifier**

Table 112: Properties of each modifier.

Id	Name	SBO
OPG		

## **Product**

Table 113: Properties of each product.

Id	Name	SBO
Sink		

## **Kinetic Law**

Derived unit contains undeclared units

$$v_{36} = \text{kdegOPG} \cdot \text{OPG} \tag{81}$$

# 7.37 Reaction MCSF\_secretion\_by\_MSC

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

# **Reaction equation**

$$MSC \xrightarrow{MSC} MSC + MCSF$$
 (82)

#### Reactant

Table 114: Properties of each reactant.

Id	Name	SBO
MSC		

### **Modifier**

Table 115: Properties of each modifier.

Id	Name	SBO
MSC		

|--|

## **Products**

Table 116: Properties of each product.

Id	Name	SBO
MSC		
MCSF		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{37} = \text{ksecMCSFbyMSC} \cdot \text{MSC}$$
 (83)

# **7.38 Reaction** MCSF\_secretion\_by\_Ob\_pro

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

# **Reaction equation**

$$Ob\_pro \xrightarrow{Ob\_pro} Ob\_pro + MCSF$$
 (84)

#### Reactant

Table 117: Properties of each reactant.

Id	Name	SBO
Ob_pro		

#### **Modifier**

Table 118: Properties of each modifier.

Id	Name	SBO
Ob_pro		

# **Products**

Table 119: Properties of each product.

Name	SBO
	Name

**Derived unit** contains undeclared units

$$v_{38} = \text{ksecMCSFbyObpro} \cdot \text{Ob\_pro}$$
 (85)

# **7.39 Reaction** MCSF\_secretion\_by\_Ob\_p

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

# **Reaction equation**

$$Ob_{-p} \xrightarrow{Ob_{-p}} Ob_{-p} + MCSF$$
 (86)

## Reactant

Table 120: Properties of each reactant.

Id	Name	SBO
0b_p		

## **Modifier**

Table 121: Properties of each modifier.

Id	Name	SBO
0b_p		

# **Products**

Table 122: Properties of each product.

Id	Name	SBO
Ob_p MCSF		

**Derived unit** contains undeclared units

$$v_{39} = \text{ksecMCSFbyObp} \cdot \text{Ob}_{-p} \tag{87}$$

# 7.40 Reaction MCSF\_secretion\_by\_Ob\_p\_PTH

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

# **Reaction equation**

$$Ob\_p\_PTH \xrightarrow{Ob\_p\_PTH} Ob\_p\_PTH + MCSF$$
 (88)

#### Reactant

Table 123: Properties of each reactant.

Id	Name	SBO
Ob_p_PTH		

#### **Modifier**

Table 124: Properties of each modifier.

Id	Name	SBO
Ob_p_PTH		

## **Products**

Table 125: Properties of each product.

Id	Name	SBO
Ob_p_PTH MCSF		

## **Kinetic Law**

Derived unit contains undeclared units

$$v_{40} = \text{ksecMCSFbyObp} \cdot \text{Ob\_p\_PTH}$$
 (89)

# 7.41 Reaction MCSF\_secretion\_by\_Ob\_m

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

# **Reaction equation**

$$Ob\_m \xrightarrow{Ob\_m} Ob\_m + MCSF$$
 (90)

#### Reactant

Table 126: Properties of each reactant.

Id	Name	SBO
Ob_m		

#### **Modifier**

Table 127: Properties of each modifier.

Id	Name	SBO
Ob_m		

#### **Products**

Table 128: Properties of each product.

Id	Name	SBO
Ob_m		
MCSF		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{41} = \text{ksecMCSFbyObm} \cdot \text{Ob}_{\text{m}}$$
 (91)

# 7.42 Reaction MCSF\_secretion\_by\_Ob\_m\_PTH

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

## **Reaction equation**

$$Ob\_m\_PTH \xrightarrow{Ob\_m\_PTH} Ob\_m\_PTH + MCSF$$
 (92)

## Reactant

Table 129: Properties of each reactant.

Id	Name	SBO
Ob_m_PTH		

## **Modifier**

Table 130: Properties of each modifier.

Id	Name	SBO
Ob_m_PTH		

## **Products**

Table 131: Properties of each product.

Id	Name	SBO
Ob_m_PTH		
MCSF		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{42} = \text{ksecMCSFbyObm} \cdot \text{Ob\_m\_PTH}$$
 (93)

# 7.43 Reaction MCSF\_degradation

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

## **Reaction equation**

$$MCSF \xrightarrow{MCSF} Sink$$
 (94)

# Reactant

Table 132: Properties of each reactant.

Id	Name	SBO
MCSF		

## **Modifier**

Table 133: Properties of each modifier.

Id	Name	SBO
MCSF		

# **Product**

Table 134: Properties of each product.

Id	Name	SBO
Sink		

## **Kinetic Law**

Derived unit contains undeclared units

$$v_{43} = \text{kdegMCSF} \cdot \text{MCSF} \tag{95}$$

# **7.44 Reaction** OPG\_secretion\_by\_Ob\_p

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

# **Reaction equation**

$$Ob_{-p} \xrightarrow{Ob_{-p}} Ob_{-p} + OPG$$
 (96)

# Reactant

Table 135: Properties of each reactant.

Id	Name	SBO
Ob_p		

### Modifier

Table 136: Properties of each modifier.

Id	Name	SBO
0b_p		

# **Products**

Table 137: Properties of each product.

Id	Name	SBO
Ob_p OPG		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{44} = ksecOPGbyObp \cdot Ob_p$$
 (97)

# 7.45 Reaction OPG\_secretion\_by\_Ob\_p\_PTH

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

# **Reaction equation**

$$Ob\_p\_PTH \xrightarrow{Ob\_p\_PTH} Ob\_p\_PTH + OPG$$

$$(98)$$

# Reactant

Table 138: Properties of each reactant.

Id	Name	SBO
Ob_p_PTH		

#### **Modifier**

Table 139: Properties of each modifier.

Id	Name	SBO
Ob_p_PTH		

## **Products**

Table 140: Properties of each product.

Id	Name	SBO
Ob_p_PTH OPG		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{45} = \text{ksecOPGbyObpPTH} \cdot \text{Ob\_p\_PTH}$$
 (99)

# 7.46 Reaction OPG\_secretion\_by\_Ob\_m

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

# **Reaction equation**

$$Ob\_m \xrightarrow{Ob\_m} Ob\_m + OPG$$
 (100)

#### Reactant

Table 141: Properties of each reactant.

Id	Name	SBO
Ob_m		

#### **Modifier**

Table 142: Properties of each modifier.

Id	Name	SBO
$\tt Ob\_m$		

# **Products**

Table 143: Properties of each product.

Id	Name	SBO
Ob_m OPG		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{46} = ksecOPGbyObm \cdot Ob_m$$
 (101)

# 7.47 Reaction RANKL\_secretion\_by\_Ocy\_A

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

# **Reaction equation**

$$Ocy_A \xrightarrow{Ocy_A} Ocy_A + RANKL$$
 (102)

#### Reactant

Table 144: Properties of each reactant.

Id	Name	SBO
Ocy_A		

# **Modifier**

Table 145: Properties of each modifier.

Id	Name	SBO
Ocy_A		

#### **Products**

Table 146: Properties of each product.

Id	Name	SBO
Ocy_A		
RANKL		

**Derived unit** contains undeclared units

$$v_{47} = ksecRANKLbyOcy \cdot Ocy\_A$$
 (103)

# 7.48 Reaction RANKL\_secretion\_by\_Ocy\_I

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

# **Reaction equation**

$$Ocy_{-}I \xrightarrow{Ocy_{-}I} Ocy_{-}I + RANKL$$
 (104)

## Reactant

Table 147: Properties of each reactant.

Id	Name	SBO
Ocy_I		

#### **Modifier**

Table 148: Properties of each modifier.

Id	Name	SBO
Ocy_I		

# **Products**

Table 149: Properties of each product.

Id	Name	SBO
Ocy_I		
RANKL		

**Derived unit** contains undeclared units

$$v_{48} = ksecRANKLbyOcyI \cdot Ocy_I$$
 (105)

# 7.49 Reaction RANKL\_secretion\_by\_MSCs

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

# **Reaction equation**

$$MSC \xrightarrow{MSC} MSC + RANKL$$
 (106)

#### Reactant

Table 150: Properties of each reactant.

Id	Name	SBO
MSC		

### **Modifier**

Table 151: Properties of each modifier.

Id	Name	SBO
MSC		

#### **Products**

Table 152: Properties of each product.

Id	Name	SBO
MSC		
RANKL		

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{49} = \text{ksecRANKLbyMSC} \cdot \text{MSC} \tag{107}$$

# **7.50 Reaction** RANKL\_secretion\_by\_Ob\_p

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

# **Reaction equation**

$$Ob_{-p} \xrightarrow{Ob_{-p}} Ob_{-p} + RANKL$$
 (108)

#### Reactant

Table 153: Properties of each reactant.

Id	Name	SBO
0b_p		

## **Modifier**

Table 154: Properties of each modifier.

Id	Name	SBO
Ob_p		

## **Products**

Table 155: Properties of each product.

Id	Name	SBO
Ob_p		
RANKL		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{50} = ksecRANKLbyObp \cdot Ob_p$$
 (109)

# 7.51 Reaction RANKL\_secretion\_by\_Ob\_p\_Tgfb\_A

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

## **Reaction equation**

$$Ob\_p\_Tgfb\_A \xrightarrow{Ob\_p\_Tgfb\_A} Ob\_p\_Tgfb\_A + RANKL$$
 (110)

#### Reactant

Table 156: Properties of each reactant.

Id	Name	SBO
Ob_p_Tgfb_A		

## **Modifier**

Table 157: Properties of each modifier.

Id	Name	SBO
Ob_p_Tgfb_A		

## **Products**

Table 158: Properties of each product.

Id	Name	SBO
Ob_p_Tgfb_A RANKL		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{51} = \text{ksecRANKLbyObpTgfb} \cdot \text{Ob\_p\_Tgfb\_A}$$
 (111)

# 7.52 Reaction RANKL\_secretion\_by\_Ob\_p\_PTH

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

# **Reaction equation**

$$Ob\_p\_PTH \xrightarrow{Ob\_p\_PTH} Ob\_p\_PTH + RANKL$$
 (112)

#### Reactant

Table 159: Properties of each reactant.

Id	Name	SBO
Ob_p_PTH		

## **Modifier**

Table 160: Properties of each modifier.

Id	Name	SBO
Ob_p_PTH		

## **Products**

Table 161: Properties of each product.

Id	Name	SBO
Ob_p_PTH		_
RANKL		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{52} = \text{ksecRANKLbyObpPTH} \cdot \text{Ob\_p\_PTH}$$
 (113)

# 7.53 Reaction RANKL\_secretion\_by\_Ob\_pro

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

# **Reaction equation**

$$Ob\_pro \xrightarrow{Ob\_pro} Ob\_pro + RANKL$$
 (114)

# Reactant

Table 162: Properties of each reactant.

Id	Name	SBO
Ob_pro		

### Modifier

Table 163: Properties of each modifier.

Id	Name	SBO
Ob_pro		

## **Products**

Table 164: Properties of each product.

Id	Name	SBO
Ob_pro		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{53} = \text{ksecRANKLbyObpro} \cdot \text{Ob\_pro}$$
 (115)

# 7.54 Reaction RANKL\_secretion\_by\_Ob\_m

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

# **Reaction equation**

$$Ob\_m \xrightarrow{Ob\_m} Ob\_m + RANKL$$
 (116)

#### Reactant

Table 165: Properties of each reactant.

Id	Name	SBO
Ob_m		

## **Modifier**

Table 166: Properties of each modifier.

Id	Name	SBO
Ob_m		

## **Products**

Table 167: Properties of each product.

Id	Name	SBO
Ob_m		
RANKL		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{54} = ksecRANKLbyObm \cdot Ob_m$$
 (117)

# 7.55 Reaction RANKL\_secretion\_by\_Ob\_m\_PTH\_enhanced

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

# **Reaction equation**

$$Ob\_m\_PTH \xrightarrow{Ob\_m\_PTH} Ob\_m\_PTH + RANKL$$
 (118)

#### Reactant

Table 168: Properties of each reactant.

Id	Name	SBO
Ob_m_PTH		

#### **Modifier**

Table 169: Properties of each modifier.

Id	Name	SBO
Ob_m_PTH		

# **Products**

Table 170: Properties of each product.

Id	Name	SBO
Ob_m_PTH		
RANKL		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{55} = \text{ksecRANKLbyObmPTH} \cdot \text{Ob\_m\_PTH}$$
 (119)

# **7.56 Reaction** Tgfb\_secretion\_by\_Obm

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

# **Reaction equation**

$$Ob\_m \xrightarrow{Ob\_m} Ob\_m + Tgfb\_I$$
 (120)

# Reactant

Table 171: Properties of each reactant.

Id	Name	SBO
Ob_m		

## Modifier

Table 172: Properties of each modifier.

Id	Name	SBO
Ob_m		

## **Products**

Table 173: Properties of each product.

Id	Name	SBO
Ob_m		
Tgfb_I		

**Derived unit** contains undeclared units

$$v_{56} = ksecTgfb \cdot Ob_m$$
 (121)

# 7.57 Reaction Tgfb\_activation

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

# **Reaction equation**

$$Tgfb\_I + Ocl\_m \xrightarrow{Tgfb\_I, Ocl\_m} Tgfb\_A + Ocl\_m$$
 (122)

## **Reactants**

Table 174: Properties of each reactant.

Id	Name	SBO
Tgfb_I Ocl_m		

### **Modifiers**

Table 175: Properties of each modifier.

Id	Name	SBO
Tgfb_I Ocl_m		

### **Products**

Table 176: Properties of each product.

Name	SBO
	Name

**Derived unit** contains undeclared units

$$v_{57} = \text{kactTgfb} \cdot \text{Tgfb} \cdot \text{I} \cdot \text{Ocl} \cdot \text{m}$$
 (123)

# 7.58 Reaction Tgfb\_degradation

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

# **Reaction equation**

$$Tgfb\_A \xrightarrow{Tgfb\_A} Sink$$
 (124)

#### Reactant

Table 177: Properties of each reactant.

Id	Name	SBO
Tgfb_A		

## **Modifier**

Table 178: Properties of each modifier.

Id	Name	SBO
Tgfb_A		

# **Product**

Table 179: Properties of each product.

Id	Name	SBO
Sink		

**Derived unit** contains undeclared units

$$v_{58} = \text{kdegTgfb} \cdot \text{Tgfb\_A} \tag{125}$$

# 7.59 Reaction Tgfb\_degradation\_by\_PTH

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

# **Reaction equation**

$$Tgfb\_A + Ob\_m\_PTH \xrightarrow{Tgfb\_A, Ob\_m\_PTH} Ob\_m\_PTH$$
 (126)

#### **Reactants**

Table 180: Properties of each reactant.

Id	Name	SBO
Tgfb_A Ob m PTH		
UD_III_PIH		

#### **Modifiers**

Table 181: Properties of each modifier.

Id	Name	SBO
Tgfb_A Ob_m_PTH		

## **Product**

Table 182: Properties of each product.

Id	Name	SBO
Ob_m_PTH		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{59} = \text{kdegTgfbPTH} \cdot \text{Tgfb\_A} \cdot \text{Ob\_m\_PTH}$$
 (127)

# 7.60 Reaction PTH\_production

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

# **Reaction equation**

Source 
$$\xrightarrow{\text{Source}}$$
 PTH (128)

#### Reactant

Table 183: Properties of each reactant.

Id	Name	SBO
Source		

#### **Modifier**

Table 184: Properties of each modifier.

Id	Name	SBO
Source		

#### **Product**

Table 185: Properties of each product.

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{60} = \text{ksynPTH} \cdot \text{Source}$$
 (129)

# 7.61 Reaction PTH\_degradation

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

# **Reaction equation**

$$PTH \xrightarrow{PTH} Sink \tag{130}$$

## Reactant

Table 186: Properties of each reactant.

Id	Name	SBO
PTH		

## **Modifier**

Table 187: Properties of each modifier.

Id	Name	SBO
PTH		

## **Product**

Table 188: Properties of each product.

Id	Name	SBO
Sink		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{61} = kdegPTH \cdot PTH \tag{131}$$

## 7.62 Reaction CREB\_activation

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

# **Reaction equation**

$$Ob\_m\_PTH + CREB \xrightarrow{CREB, Ob\_m\_PTH} Ob\_m\_PTH + CREB\_P$$
 (132)

## **Reactants**

Table 189: Properties of each reactant.

Id	Name	SBO
Ob_m_PTH		
CREB		

## **Modifiers**

Table 190: Properties of each modifier.

Id	Name	SBO
CREB		
${\tt Ob\_m\_PTH}$		

## **Products**

Table 191: Properties of each product.

Id	Name	SBO
Ob_m_PTH		
$CREB_P$		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{62} = \frac{\text{kactCreb} \cdot \text{CREB} \cdot \text{Ob\_m\_PTH}^2}{100^2 + \text{Ob\_m\_PTH}^2}$$
(133)

## 7.63 Reaction CREB\_inactivation

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

## **Reaction equation**

$$CREB\_P \xrightarrow{CREB\_P} CREB \tag{134}$$

## Reactant

Table 192: Properties of each reactant.

Id	Name	SBO
CREB_P		

## **Modifier**

Table 193: Properties of each modifier.

Id	Name	SBO
CREB_P		

## **Product**

Table 194: Properties of each product.

Id	Name	SBO
CREB		

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{63} = \text{kinactCreb} \cdot \text{CREB\_P}$$
 (135)

# 7.64 Reaction CREB\_Runx2\_binding

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

# **Reaction equation**

$$CREB\_P + Runx2 \xrightarrow{CREB\_P, Runx2} CREB\_Runx2$$
 (136)

#### **Reactants**

Table 195: Properties of each reactant.

Id	Name	SBO
CREB_P		
Runx2		

# **Modifiers**

Table 196: Properties of each modifier.

Id	Name	SBO
CREB_P		
Runx2		

### **Product**

Table 197: Properties of each product.

Id	Name	SBO
CREB_Runx2		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{64} = \text{kbinCrebRunx2} \cdot \text{CREB\_P} \cdot \text{Runx2}$$
 (137)

# 7.65 Reaction CREB\_Runx2\_release

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

# **Reaction equation**

$$CREB\_Runx2 \xrightarrow{CREB\_Runx2} CREB\_P + Runx2$$
 (138)

### Reactant

Table 198: Properties of each reactant.

Id	Name	SBO
CREB_Runx2		

# **Modifier**

Table 199: Properties of each modifier.

Id	Name	SBO
CREB_Runx2		

# **Products**

Table 200: Properties of each product.

Id	Name	SBO
CREB_P		
Runx2		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{65} = \text{krelCrebRunx2} \cdot \text{CREB\_Runx2}$$
 (139)

# 7.66 Reaction Bcl2\_synthesis

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

# **Reaction equation**

$$CREB\_Runx2 \xrightarrow{CREB\_Runx2} CREB\_Runx2 + Bcl2$$
 (140)

### Reactant

Table 201: Properties of each reactant.

Id	Name	SBO
CREB_Runx2		

### **Modifier**

74

Table 202: Properties of each modifier.

Id	Name	SBO
CREB_Runx2		

# **Products**

Table 203: Properties of each product.

Id	Name	SBO
CREB_Runx2		
Bcl2		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{66} = \text{ksynBcl2} \cdot \text{CREB\_Runx2}$$
 (141)

# 7.67 Reaction Bcl2\_degradation

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

# **Reaction equation**

$$Bcl2 \xrightarrow{Bcl2} Sink \tag{142}$$

# Reactant

Table 204: Properties of each reactant.

Id	Name	SBO
Bcl2		

# Modifier

Table 205: Properties of each modifier.

Id	Name	SBO
Bcl2		

# **Product**

Table 206: Properties of each product.

Id	Name	SBO
Sink		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{67} = \text{kdegBcl2} \cdot \text{Bcl2} \tag{143}$$

# 7.68 Reaction Bax\_Bcl2\_binding

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

# **Reaction equation**

$$Bax + Bcl2 \xrightarrow{Bax, Bcl2} Bax\_Bcl2$$
 (144)

### **Reactants**

Table 207: Properties of each reactant.

Id	Name	SBO
Bax		
Bc12		

### **Modifiers**

Table 208: Properties of each modifier.

Id	Name	SBO
Bax		
Bcl2		

# **Product**

Table 209: Properties of each product.

Id	Name	SBO
Bax_Bc12		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{68} = \text{kbinBaxBcl2} \cdot \text{Bax} \cdot \text{Bcl2} \tag{145}$$

# 7.69 Reaction Bax\_Bcl2\_release

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

# **Reaction equation**

$$Bax\_Bcl2 \xrightarrow{Bax\_Bcl2} Bax + Bcl2$$
 (146)

### Reactant

Table 210: Properties of each reactant.

Id	Name	SBO
Bax_Bc12		

# **Modifier**

Table 211: Properties of each modifier.

Id	Name	SBO
Bax_Bcl2		

### **Products**

Table 212: Properties of each product.

Id	Name	SBO
Bax		
Bcl2		

### **Kinetic Law**

Derived unit contains undeclared units

$$v_{69} = \text{krelBaxBcl2} \cdot \text{Bax\_Bcl2} \tag{147}$$

# 7.70 Reaction Runx2\_synthesis

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

# **Reaction equation**

Source 
$$\xrightarrow{\text{Source}} \text{Runx2}$$
 (148)

#### Reactant

Table 213: Properties of each reactant.

Id	Name	SBO
Source		

### **Modifier**

Table 214: Properties of each modifier.

Id	Name	SBO
Source		

### **Product**

Table 215: Properties of each product.

Id	Name	SBO
Runx2		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{70} = \text{ksynRunx2} \cdot \text{Source}$$
 (149)

# 7.71 Reaction Runx2\_degradation\_via\_PTH

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

# **Reaction equation**

$$Ob\_m\_PTH + Runx2 \xrightarrow{Runx2, Ob\_m\_PTH} Ob\_m\_PTH$$
 (150)

### **Reactants**

Table 216: Properties of each reactant.

Id	Name	SBO
Ob_m_PTH		
Runx2		

### **Modifiers**

Table 217: Properties of each modifier.

Id	Name	SBO
Runx2		
$\texttt{Ob\_m\_PTH}$		

# **Product**

Table 218: Properties of each product.

Id	Name	SBO
Ob_m_PTH		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{71} = \text{kdegRunx2PTH} \cdot \text{Runx2} \cdot \text{Ob\_m\_PTH}$$
 (151)

# 7.72 Reaction Runx2\_degradation

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

# **Reaction equation**

$$Runx2 \xrightarrow{Runx2} Sink$$
 (152)

# Reactant

Table 219: Properties of each reactant.

Id	Name	SBO
Runx2		

# **Modifier**

Table 220: Properties of each modifier.

Id	Name	SBO
Runx2		

# **Product**

Table 221: Properties of each product.

Id	Name	SBO
Sink		

# **Kinetic Law**

Derived unit contains undeclared units

$$v_{72} = kdegRunx2 \cdot Runx2 \tag{153}$$

# 7.73 Reaction Bone\_formation

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

# **Reaction equation**

$$Ob_{-m} \xrightarrow{Ob_{-m}} Ob_{-m} + Bone + newbone$$
 (154)

### Reactant

Table 222: Properties of each reactant.

Id	Name	SBO
Ob_m		

### **Modifier**

Table 223: Properties of each modifier.

Id	Name	SBO
Ob_m		

# **Products**

Table 224: Properties of each product.

Id	Name	SBO
Ob_m		
Bone		
newbone		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{73} = \text{kformBone} \cdot \text{Ob}_{\text{-m}}$$
 (155)

# 7.74 Reaction Bone\_formation\_Obm\_PTH

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

# **Reaction equation**

$$Ob\_m\_PTH \xrightarrow{Ob\_m\_PTH} Ob\_m\_PTH + Bone + newbone$$
 (156)

# Reactant

Table 225: Properties of each reactant.

Id	Name	SBO
Ob_m_PTH		

# **Modifier**

Table 226: Properties of each modifier.

Id	Name	SBO
Ob_m_PTH		

# **Products**

Table 227: Properties of each product.

Id	Name	SBO
Ob_m_PTH		
Bone		
newbone		

# **Kinetic Law**

Derived unit contains undeclared units

$$v_{74} = \text{kformBone} \cdot \text{Ob\_m\_PTH}$$
 (157)

# 7.75 Reaction Bone\_degradation

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

# **Reaction equation**

$$Ocl_{-m} + Bone \xrightarrow{Ocl_{-m}, Bone} Ocl_{-m}$$
 (158)

### **Reactants**

Table 228: Properties of each reactant.

Id	Name	SBO
Ocl_m		
Bone		

# **Modifiers**

Table 229: Properties of each modifier.

Id	Name	SBO
Ocl_m		
Bone		

# **Product**

Table 230: Properties of each product.

Id	Name	SBO
Ocl_m		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{75} = \text{kdegBone} \cdot \text{Ocl}_{-m} \cdot \text{Bone}$$
 (159)

# 7.76 Reaction Xsynthesis

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

# **Reaction equation**

Source 
$$\xrightarrow{\text{Source}} X$$
 (160)

### Reactant

Table 231: Properties of each reactant.

Id	Name	SBO
Source		

### **Modifier**

Table 232: Properties of each modifier.

Id	Name	SBO
Source		

### **Product**

Table 233: Properties of each product.

Id	Name	SBO
X		

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{76} = \text{ksynX} \cdot \text{Source}$$
 (161)

# 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 Bone

### Initial amount 2000 item

This species takes part in four reactions (as a reactant in Bone\_degradation and as a product in Bone\_formation, Bone\_formation\_Obm\_PTH and as a modifier in Bone\_degradation).

$$\frac{d}{dt}Bone = |v_{73}| + |v_{74}| - |v_{75}| \tag{162}$$

### 8.2 Species HSC

### Initial amount 100 item

This species takes part in three reactions (as a reactant in HSC\_differentiation\_to\_Ocl\_p and as a product in HSC\_differentiation\_to\_Ocl\_p and as a modifier in HSC\_differentiation\_to\_Ocl\_p).

$$\frac{d}{dt}HSC = |v_{26}| - |v_{26}| \tag{163}$$

### 8.3 Species Ob\_m

### **Initial amount** 0 item

This species takes part in 25 reactions (as a reactant in Ob\_maturation\_to\_Ocy, Ob\_maturation\_to\_Ocy\_by\_Tgfb, Ob\_m\_bound\_by\_PTH, Ob\_m\_apoptosis, MCSF\_secretion\_by\_Ob\_m, OPG\_secretion\_by\_Ob\_m, RANKL\_secretion\_by\_Ob\_m, Tgfb\_secretion\_by\_Obm, Bone\_formation and as a product in Ob\_precursor\_differentiation, Ob\_m\_PTH\_release, MCSF\_secretion\_by\_Ob\_m, OPG\_secretion\_by\_Ob\_m, RANKL\_secretion\_by\_Ob\_m, Tgfb\_secretion\_by\_Obm, Bone\_formation and as a modifier in Ob\_maturation\_to\_Ocy, Ob\_maturation\_to\_Ocy\_by\_Tgfb, Ob\_m\_bound\_by\_PTH, Ob\_m\_apoptosis, MCSF\_secretion\_by\_Ob\_m, OPG\_secretion\_by\_Ob\_m, RANKL\_secretion\_by\_Ob\_m, Tgfb\_secretion\_by\_Ob\_m, Bone\_formation).

$$\frac{d}{dt}Ob_{m} = v_{15} + v_{23} + v_{41} + v_{46} + v_{54} + v_{56} + v_{73} - v_{20}$$

$$- v_{21} - v_{22} - v_{24} - v_{41} - v_{46} - v_{54} - v_{56} - v_{73}$$

$$(164)$$

# 8.4 Species Ob\_p

#### **Initial amount** 0 item

This species takes part in 18 reactions (as a reactant in Ob\_precursor\_differentiation, Ob\_p\_binding\_by\_PTH, Ob\_p\_Tgfb\_binding, MCSF\_secretion\_by\_Ob\_p, OPG\_secretion\_by\_Ob\_p, CDb\_p, RANKL\_secretion\_by\_Ob\_p and as a product in Osteoblast\_progenitor\_differentiation\_by\_Tgfb, Ob\_p\_PTH\_release, Ob\_p\_Tgfb\_release, MCSF\_secretion\_by\_Ob\_p, OPG\_secretion\_by\_Ob\_p, RANKL\_secretion\_by\_Ob\_p and as a modifier in Ob\_precursor\_differentiation, Ob\_p\_binding\_by\_PTH, Ob\_p\_Tgfb\_binding, MCSF\_secretion\_by\_Ob\_p, OPG\_secretion\_by\_Ob\_p, RANKL\_secretion\_by\_Ob\_p).

$$\frac{d}{dt}Ob_{-}p = v_{14} + v_{17} + v_{19} + v_{39} + v_{44} + v_{50} - v_{15} - v_{16} - v_{18} - v_{39} - v_{44} - v_{50}$$
(165)

# 8.5 Species Ob\_p\_Tgfb\_A

#### **Initial amount** 0 item

This species takes part in six reactions (as a reactant in Ob\_p\_Tgfb\_release, RANKL\_secretion\_by\_Ob\_p\_Tgfb\_A and as a product in Ob\_p\_Tgfb\_binding, RANKL\_secretion\_by\_Ob\_p\_Tgfb\_A and as a modifier in Ob\_p\_Tgfb\_release, RANKL\_secretion\_by\_Ob\_p\_Tgfb\_A).

$$\frac{d}{dt}Ob_{-}p_{-}Tgfb_{-}A = |v_{18}| + |v_{51}| - |v_{19}| - |v_{51}|$$
(166)

### 8.6 Species Ob\_pro

#### **Initial amount** 0 item

This species takes part in nine reactions (as a reactant in Osteoblast\_progenitor\_differentiation\_by\_Tgfb, MCSF\_secretion\_by\_Ob\_pro, RANKL\_secretion\_by\_Ob\_pro and as a product in MSC\_differentiation\_to\_Ob\_pro, MCSF\_secretion\_by\_Ob\_pro, RANKL\_secretion\_by\_Ob\_pro and as a modifier in Osteoblast\_progenitor\_differentiation\_by\_Tgfb, MCSF\_secretion\_by\_Ob\_pro, RANKL\_secretion\_by\_Ob\_pro).

$$\frac{d}{dt}Ob\_pro = |v_{13}| + |v_{38}| + |v_{53}| - |v_{14}| - |v_{38}| - |v_{53}|$$
(167)

### 8.7 Species Ocl\_m

### **Initial amount** 0 item

This species takes part in nine reactions (as a reactant in Osteoclast\_apoptosis, Tgfb-activation, Bone\_degradation and as a product in Osteoclast\_precursor\_differentiation, Tgfb\_activation, Bone\_degradation and as a modifier in Osteoclast\_apoptosis, Tgfb-activation, Bone\_degradation).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Ocl}_{m} = v_{30} + v_{57} + v_{75} - v_{34} - v_{57} - v_{75} \tag{168}$$

### 8.8 Species Ocl\_p

#### **Initial amount** 0 item

This species takes part in six reactions (as a reactant in Ocl\_p\_apoptosis, Ocl\_p\_RANKL\_binding and as a product in HSC\_differentiation\_to\_Ocl\_p, Ocl\_p\_RANKL\_release and as a modifier in Ocl\_p\_apoptosis, Ocl\_p\_RANKL\_binding).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Ocl}_{-p} = |v_{26}| + |v_{29}| - |v_{27}| - |v_{28}| \tag{169}$$

### 8.9 Species Ocl\_p\_RANKL

#### Initial amount 0 item

This species takes part in five reactions (as a reactant in Ocl\_p\_RANKL\_release, Osteoclast\_precursor\_differentiation and as a product in Ocl\_p\_RANKL\_binding and as a modifier in Ocl\_p\_RANKL\_release, Osteoclast\_precursor\_differentiation).

$$\frac{d}{dt} \text{Ocl\_p\_RANKL} = |v_{28}| - |v_{29}| - |v_{30}|$$
 (170)

# 8.10 Species Ocy\_A

#### **Initial amount** 0 item

This species takes part in seven reactions (as a reactant in Osteocyte\_inactivation, RANKL-secretion\_by\_Ocy\_A and as a product in Osteocyte\_activation, Osteocyte\_activation-by\_PTH, RANKL\_secretion\_by\_Ocy\_A and as a modifier in Osteocyte\_inactivation, RANKL-secretion\_by\_Ocy\_A).

$$\frac{d}{dt}Ocy_A = |v_2| + |v_3| + |v_{47}| - |v_4| - |v_{47}|$$
(171)

# 8.11 Species Ocy\_I

### Initial amount 1800 item

This species takes part in 16 reactions (as a reactant in Osteocyte\_activation, Ocy\_I\_bound\_by\_PTH, Ocy\_apoptosis, Sost\_secretion, RANKL\_secretion\_by\_Ocy\_I and as a product in Osteocyte\_inactivation, Ocy\_I\_PTH\_release, Sost\_secretion, Ob\_maturation\_to\_Ocy, Ob\_maturation\_to\_Ocy\_by\_Tgfb, RANKL\_secretion\_by\_Ocy\_I and as a modifier in Osteocyte\_activation, Ocy\_I\_bound\_by\_PTH, Ocy\_apoptosis, Sost\_secretion, RANKL\_secretion\_by\_Ocy\_I).

$$\frac{d}{dt}Ocy I = v_4 + v_6 + v_8 + v_{20} + v_{21} + v_{48} - v_2 - v_5 - v_7 - v_8 - v_{48}$$
 (172)

# 8.12 Species Ocy\_I\_PTH

### **Initial amount** 0 item

This species takes part in five reactions (as a reactant in Osteocyte\_activation\_by\_PTH, Ocy\_I\_PTH\_release and as a product in Ocy\_I\_bound\_by\_PTH and as a modifier in Osteocyte\_activation\_by\_PTH, Ocy\_I\_PTH\_release).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Ocy}\mathrm{LPTH} = |v_5| - |v_3| - |v_6| \tag{173}$$

### 8.13 Species MSC

# Initial amount 100 item

This species takes part in nine reactions (as a reactant in MSC\_differentiation\_to\_Ob\_pro, MCSF\_secretion\_by\_MSC, RANKL\_secretion\_by\_MSCs and as a product in MSC\_differentiation\_to\_Ob\_pro, MCSF\_secretion\_by\_MSC, RANKL\_secretion\_by\_MSCs and as a modifier in MSC\_differentiation\_to\_Ob\_pro, MCSF\_secretion\_by\_MSC, RANKL\_secretion\_by\_MSCs).

$$\frac{d}{dt}MSC = v_{13} + v_{37} + v_{49} - v_{13} - v_{37} - v_{49}$$
(174)

# 8.14 Species LOAD

#### **Initial amount** 0 item

### Involved in events AddLOAD1, AddLOAD2

This species takes part in five reactions (as a reactant in Unloading, Osteocyte\_activation and as a product in Osteocyte\_activation and as a modifier in Unloading, Osteocyte\_activation).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{LOAD} = |v_2| - |v_1| - |v_2| \tag{175}$$

Furthermore, two events influence this species' rate of change.

### 8.15 Species MCSF

#### **Initial amount** 5 item

This species takes part in eleven reactions (as a reactant in HSC\_differentiation\_to\_Ocl\_p, MCSF\_degradation and as a product in HSC\_differentiation\_to\_Ocl\_p, MCSF\_secretion\_by\_MSC, MCSF\_secretion\_by\_Ob\_pro, MCSF\_secretion\_by\_Ob\_p, MCSF\_secretion\_by\_Ob\_p\_PTH, MCSF\_secretion\_by\_Ob\_m, MCSF\_secretion\_by\_Ob\_m\_PTH and as a modifier in HSC\_differentiation\_to\_Ocl\_p, MCSF\_degradation).

$$\frac{d}{dt}MCSF = v_{26} + v_{37} + v_{38} + v_{39} + v_{40} + v_{41} + v_{42} - v_{26} - v_{43}$$
 (176)

### 8.16 Species Ob\_m\_PTH

### **Initial amount** 0 item

This species takes part in 26 reactions (as a reactant in Wnt\_activation\_by\_PTH, 0b\_m\_PTH\_release, 0b\_m\_PTH\_apoptosis, MCSF\_secretion\_by\_0b\_m\_PTH, RANKL\_secretion\_by\_0b\_m\_PTH\_enhanced, Tgfb\_degradation\_by\_PTH, CREB\_activation, Runx2\_degradation\_via\_PTH, Bone\_formation\_0bm\_PTH and as a product in Wnt\_activation\_by\_PTH, 0b\_m\_bound\_by\_PTH, MCSF\_secretion\_by\_0b\_m\_PTH, RANKL\_secretion\_by\_0b\_m\_PTH\_enhanced, Tgfb\_degradation\_by\_PTH, CREB\_activation, Runx2\_degradation\_via\_PTH, Bone\_formation\_0bm\_PTH and as a modifier in Wnt\_activation\_by\_PTH, 0b\_m\_PTH\_release, 0b\_m\_PTH\_apoptosis, MCSF\_secretion\_by\_0b\_m\_PTH, RANKL\_secretion\_by\_0b\_m\_PTH\_enhanced, Tgfb\_degradation\_by\_PTH, CREB\_activation, Runx2\_degradation\_via\_PTH, Bone\_formation\_0bm\_PTH).

$$\frac{d}{dt}Ob\_m\_PTH = v_{11} + v_{22} + v_{42} + v_{55} + v_{59} + v_{62} + v_{71} + v_{74} - v_{11} 
- v_{23} - v_{25} - v_{42} - v_{55} - v_{59} - v_{62} - v_{71} - v_{74}$$
(177)

# 8.17 Species Ob\_p\_PTH

#### **Initial amount** 0 item

This species takes part in twelve reactions (as a reactant in Ob\_p\_PTH\_release, MCSF\_secretion\_by\_Ob\_p\_PTH, OPG\_secretion\_by\_Ob\_p\_PTH, RANKL\_secretion\_by\_Ob\_p\_PTH and as a product in Ob\_p\_binding\_by\_PTH, MCSF\_secretion\_by\_Ob\_p\_PTH, OPG\_secretion\_by\_Ob\_p\_PTH, RANKL\_secretion\_by\_Ob\_p\_PTH and as a modifier in Ob\_p\_PTH\_release, MCSF\_secretion\_by\_Ob\_p\_PTH, OPG\_secretion\_by\_Ob\_p\_PTH, RANKL\_secretion\_by\_Ob\_p\_PTH, RANKL\_secretion\_by\_Ob\_p\_PTH).

$$\frac{d}{dt}Ob_{-}p_{-}PTH = v_{16} + v_{40} + v_{45} + v_{52} - v_{17} - v_{40} - v_{45} - v_{52}$$
 (178)

### 8.18 Species OPG

### **Initial amount** 0 item

This species takes part in eight reactions (as a reactant in RANKL\_inhibition, OPG\_degradation and as a product in OPG\_RANKL\_dissociation, OPG\_secretion\_by\_Ob\_p, OPG\_secretion\_by\_Ob\_p\_PTH, OPG\_secretion\_by\_Ob\_m and as a modifier in RANKL\_inhibition, OPG\_degradation).

$$\frac{\mathrm{d}}{\mathrm{d}t} OPG = |v_{32}| + |v_{44}| + |v_{45}| + |v_{46}| - |v_{31}| - |v_{36}|$$
(179)

# 8.19 Species OPG\_RANKL

### **Initial amount** 0 item

This species takes part in five reactions (as a reactant in OPG\_RANKL\_dissociation, OPG\_RANKL\_degradation and as a product in RANKL\_inhibition and as a modifier in OPG\_RANKL\_dissociation, OPG\_RANKL\_degradation).

$$\frac{d}{dt}OPG\_RANKL = |v_{31}| - |v_{32}| - |v_{33}|$$
 (180)

# 8.20 Species PTH

### Initial amount 170 item

### Involved in events AddPTH1, AddPTH2

This species takes part in 14 reactions (as a reactant in <code>Ocy\_I\_bound\_by\_PTH</code>, <code>Ob\_p\_binding\_by\_PTH</code>, <code>Ob\_m\_bound\_by\_PTH</code>, <code>PTH\_degradation</code> and as a product in <code>Osteocyte\_activation\_by\_PTH</code>, <code>Ocy\_I\_PTH\_release</code>, <code>Ob\_m\_PTH\_release</code>, <code>Ob\_m\_PTH\_release</code>, <code>Ob\_m\_PTH\_apoptosis</code>, <code>PTH\_production</code> and as a modifier in <code>Ocy\_I\_bound\_by\_PTH</code>, <code>Ob\_p\_binding\_by\_PTH</code>, <code>Ob\_m\_bound\_by\_PTH</code>, <code>PTH\_degradation</code>).

$$\frac{d}{dt}PTH = v_3 + v_6 + v_{17} + v_{23} + v_{25} + v_{60} - v_5 - v_{16} - v_{22} - v_{61}$$
 (181)

Furthermore, two events influence this species' rate of change.

# 8.21 Species RANKL

#### **Initial amount** 0 item

This species takes part in 17 reactions (as a reactant in Ocl\_p\_RANKL\_binding, RANKL\_inhibition, RANKL\_degradation and as a product in Ocl\_p\_RANKL\_release, OPG\_RANKL\_dissociation, RANKL\_secretion\_by\_Ocy\_A, RANKL\_secretion\_by\_Ocy\_I, RANKL\_secretion\_by\_MSCs, RANKL\_secretion\_by\_Ob\_p, RANKL\_secretion\_by\_Ob\_p\_Tgfb\_A, RANKL\_secretion\_by\_Ob\_p\_PTH, RANKL\_secretion\_by\_Ob\_pro, RANKL\_secretion\_by\_Ob\_m, RANKL\_secretion\_by\_Ob\_m\_PTH-enhanced and as a modifier in Ocl\_p\_RANKL\_binding, RANKL\_inhibition, RANKL\_degradation).

$$\frac{d}{dt}RANKL = |v_{29}| + |v_{32}| + |v_{47}| + |v_{48}| + |v_{49}| + |v_{50}| + |v_{51}| + |v_{52}| + |v_{53}| + |v_{54}| + |v_{55}| - |v_{28}| - |v_{31}| - |v_{35}|$$
(182)

# 8.22 Species Sost

#### **Initial amount** 0 item

This species takes part in six reactions (as a reactant in Sost\_degradation, Inactivation\_of\_Wnt\_by\_Sost and as a product in Sost\_secretion, Inactivation\_of\_Wnt\_by\_Sost and as a modifier in Sost\_degradation, Inactivation\_of\_Wnt\_by\_Sost).

$$\frac{\mathrm{d}}{\mathrm{d}t} \text{Sost} = |v_8| + |v_{12}| - |v_9| - |v_{12}| \tag{183}$$

# 8.23 Species Tgfb\_A

#### **Initial amount** 5 item

This species takes part in 14 reactions (as a reactant in Osteoblast\_progenitor\_differentiation\_by\_Tgfb, Ob\_p\_Tgfb\_binding, Ob\_maturation\_to\_Ocy\_by\_Tgfb, Tgfb\_degradation, Tgfb\_degradation\_by\_PTH and as a product in Osteoblast\_progenitor\_differentiation\_by\_Tgfb, Ob\_p\_Tgfb\_release, Ob\_maturation\_to\_Ocy\_by\_Tgfb, Tgfb\_activation and as a modifier in Osteoblast\_progenitor\_differentiation\_by\_Tgfb, Ob\_p\_Tgfb\_binding, Ob\_maturation\_to\_Ocy\_by\_Tgfb, Tgfb\_degradation, Tgfb\_degradation\_by\_PTH).

$$\frac{d}{dt} Tgfb\_A = |v_{14}| + |v_{19}| + |v_{21}| + |v_{57}| - |v_{14}| - |v_{18}| - |v_{21}| - |v_{58}| - |v_{59}|$$
(184)

# 8.24 Species Tgfb\_I

#### Initial amount 500 item

This species takes part in three reactions (as a reactant in Tgfb\_activation and as a product in Tgfb\_secretion\_by\_Obm and as a modifier in Tgfb\_activation).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Tgfb}.\mathrm{I} = v_{56} - v_{57} \tag{185}$$

# 8.25 Species Wnt\_A

#### **Initial amount** 0 item

This species takes part in seven reactions (as a reactant in Inactivation\_of\_Wnt\_by\_Sost, MSC\_differentiation\_to\_Ob\_pro and as a product in Wnt\_activation, Wnt\_activation\_by\_PTH, MSC\_differentiation\_to\_Ob\_pro and as a modifier in Inactivation\_of\_Wnt\_by\_Sost, MSC\_differentiation\_to\_Ob\_pro).

$$\frac{d}{dt}Wnt\_A = |v_{10}| + |v_{11}| + |v_{13}| - |v_{12}| - |v_{13}|$$
(186)

### 8.26 Species Wnt\_I

#### **Initial amount** 200 item

This species takes part in five reactions (as a reactant in Wnt\_activation, Wnt\_activation—by\_PTH and as a product in Inactivation\_of\_Wnt\_by\_Sost and as a modifier in Wnt\_activation, Wnt\_activation\_by\_PTH).

$$\frac{d}{dt}Wnt_{-}I = |v_{12}| - |v_{10}| - |v_{11}|$$
(187)

### 8.27 Species Source

# **Initial amount** 1 item

This species takes part in six reactions (as a reactant in PTH\_production, Runx2\_synthesis, Xsynthesis and as a modifier in PTH\_production, Runx2\_synthesis, Xsynthesis), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Source} = 0\tag{188}$$

### 8.28 Species Sink

### **Initial amount** 1 item

This species takes part in 13 reactions (as a product in Unloading, Ocy\_apoptosis, Sost\_degradation, Ocl\_p\_apoptosis, OPG\_RANKL\_degradation, Osteoclast\_apoptosis, RANKL-degradation, OPG\_degradation, MCSF\_degradation, Tgfb\_degradation, PTH\_degradation, Bcl2\_degradation, Runx2\_degradation), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Sink} = 0\tag{189}$$

# 8.29 Species X

#### **Initial amount** 0 item

#### Involved in event AddPTH2

This species takes part in one reaction (as a product in Xsynthesis).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathbf{X} = v_{76} \tag{190}$$

Furthermore, one event influences this species' rate of change.

# 8.30 Species newbone

# **Initial amount** 0 item

This species takes part in two reactions (as a product in Bone\_formation, Bone\_formation\_\_Obm\_PTH).

$$\frac{\mathrm{d}}{\mathrm{d}t}\text{newbone} = v_{73} + v_{74} \tag{191}$$

### 8.31 Species Bax

### Initial amount 100 item

This species takes part in nine reactions (as a reactant in Ob\_m\_apoptosis, Ob\_m\_PTH\_apoptosis, Bax\_Bc12\_binding and as a product in Ob\_m\_apoptosis, Ob\_m\_PTH\_apoptosis, Bax\_Bc12\_release and as a modifier in Ob\_m\_apoptosis, Ob\_m\_PTH\_apoptosis, Bax\_Bc12\_binding).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Bax} = |v_{24}| + |v_{25}| + |v_{69}| - |v_{24}| - |v_{25}| - |v_{68}| \tag{192}$$

### 8.32 Species Bax\_Bc12

### **Initial amount** 0 item

This species takes part in three reactions (as a reactant in Bax\_Bcl2\_release and as a product in Bax\_Bcl2\_binding and as a modifier in Bax\_Bcl2\_release).

$$\frac{\mathrm{d}}{\mathrm{d}t} \mathrm{Bax}_{-} \mathrm{Bcl2} = |v_{68}| - |v_{69}| \tag{193}$$

# 8.33 Species Bc12

### **Initial amount** 0 item

This species takes part in six reactions (as a reactant in Bc12\_degradation, Bax\_Bc12\_binding and as a product in Bc12\_synthesis, Bax\_Bc12\_release and as a modifier in Bc12\_degradation, Bax\_Bc12\_binding).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Bcl2} = |v_{66}| + |v_{69}| - |v_{67}| - |v_{68}| \tag{194}$$

# 8.34 Species CREB

#### **Initial amount** 100 item

This species takes part in three reactions (as a reactant in CREB\_activation and as a product in CREB\_inactivation and as a modifier in CREB\_activation).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{CREB} = |v_{63}| - |v_{62}| \tag{195}$$

# 8.35 Species CREB\_P

#### **Initial amount** 0 item

This species takes part in six reactions (as a reactant in CREB\_inactivation, CREB\_Runx2\_binding and as a product in CREB\_activation, CREB\_Runx2\_release and as a modifier in CREB\_inactivation, CREB\_Runx2\_binding).

$$\frac{d}{dt}CREB_P = |v_{62}| + |v_{65}| - |v_{63}| - |v_{64}|$$
(196)

# 8.36 Species CREB\_Runx2

#### **Initial amount** 0 item

This species takes part in six reactions (as a reactant in CREB\_Runx2\_release, Bc12\_synthesis and as a product in CREB\_Runx2\_binding, Bc12\_synthesis and as a modifier in CREB\_Runx2\_release, Bc12\_synthesis).

$$\frac{d}{dt}CREB_Runx2 = v_{64} + v_{66} - v_{65} - v_{66}$$
 (197)

# 8.37 Species Runx2

### **Initial amount** 10 item

This species takes part in eight reactions (as a reactant in CREB\_Runx2\_binding, Runx2\_degradation\_via\_PTH, Runx2\_degradation and as a product in CREB\_Runx2\_release, Runx2\_synthesis and as a modifier in CREB\_Runx2\_binding, Runx2\_degradation\_via\_PTH, Runx2\_degradation).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Runx2} = |v_{65}| + |v_{70}| - |v_{64}| - |v_{71}| - |v_{72}| \tag{198}$$

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