## **SBML Model Report**

# Model name: "Carbo2013 - Cytokine driven CD4+ T Cell differentiation and phenotype plasticity"



May 6, 2016

### 1 General Overview

This is a document in SBML Level 2 Version 4 format. This model was created by the following two authors: Vijayalakshmi Chelliah<sup>1</sup> and Adria Carbo<sup>2</sup> at May nineth 2013 at 2:36 p.m. and last time modified at October tenth 2014 at 11:12 a.m. Table 1 gives an overview of the quantities of all components of this model.

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

Element	Quantity	Element	Quantity
compartment types	0	compartments	2
species types	0	species	94
events	0	constraints	0
reactions	53	function definitions	16
global parameters	5	unit definitions	1
rules	2	initial assignments	12

## **Model Notes**

Carbo2013 - Cytokine driven CD4+ T Cell differentiation and phenotype plasticity

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CD4+ T cells can differentiate into different phenotypes depending on the cytokine milieu. Here a computational and mathematical model with sixty ordinary differential equations representing a CD4+ T cell differentiating into either Th1, Th2, Th17 or iTreg cells, has been constructed. The model includes cytokines, nuclear receptors and transcription factors that define fate and function of CD4+ T cells. Computational simulations illustrate how a proinflammatory Th17 cell can undergo reprogramming into an anti-inflammatory iTreg phenotype following PPARc activation.

This model is described in the article: Systems Modeling of Molecular Mechanisms Controlling Cytokine-driven CD4+ T Cell Differentiation and Phenotype Plasticity. Carbo A, Hontecillas R, Kronsteiner B, Viladomiu M, Pedragosa M, Lu P, Philipson CW, Hoops S, Marathe M, Eubank S, Bisset K, Wendelsdorf K, Jarrah A, Mei Y, Bassaganya-Riera JPLoS Computational Biology [2013, 9(4):e1003027]

#### Abstract:

Differentiation of CD4+ T cells into effector or regulatory phenotypes is tightly controlled by the cytokine milieu, complex intracellular signaling networks and numerous transcriptional regulators. We combined experimental approaches and computational modeling to investigate the mechanisms controlling differentiation and plasticity of CD4+ T cells in the gut of mice. Our computational model encompasses the major intracellular pathways involved in CD4+ T cell differentiation into T helper 1 (Th1), Th2, Th17 and induced regulatory T cells (iTreg). Our modeling efforts predicted a critical role for peroxisome proliferator-activated receptor gamma (PPAR) in modulating plasticity between Th17 and iTreg cells. PPAR regulates differentiation, activation and cytokine production, thereby controlling the induction of effector and regulatory responses, and is a promising therapeutic target for dysregulated immune responses and inflammation. Our modeling efforts predict that following PPAR activation, Th17 cells undergo phenotype switch and become iTreg cells. This prediction was validated by results of adoptive transfer studies showing an increase of colonic iTreg and a decrease of Th17 cells in the gut mucosa of mice with colitis following pharmacological activation of PPAR. Deletion of PPAR in CD4+ T cells impaired mucosal iTreg and enhanced colitogenic Th17 responses in mice with CD4+ T cell-induced colitis. Thus, for the first time we provide novel molecular evidence in vivo demonstrating that PPAR in addition to regulating CD4+ T cell differentiation also plays a major role controlling Th17 and iTreg plasticity in the gut mucosa.

**Author's comment:** CD4+ T cell computational model (Version 1.4)Steady state corrected. There was a problem in the internalization of IL-17 in its mathematical function.

This model is hosted on BioModels Database and identifiedby: MODEL1304230001.

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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 time

Name time

**Definition** 3600 s

#### 2.2 Unit substance

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

**Definition** mol

#### 2.3 Unit volume

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

**Definition** 1

#### 2.4 Unit area

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

 $\textbf{Definition}\ m^2$ 

## 2.5 Unit length

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

**Definition** m

## 3 Compartments

This model contains two compartments.

Table 2: Properties of all compartments.

Id	Name	SBO	Spatial Dimensions	Size	Unit	Constant	Outside
default	default T Helper Cell	0000290 0000290	3 3	1 1	litre litre	<b>Z</b>	

## 3.1 Compartment default

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

Name default

**SBO:0000290** physical compartment

## 3.2 Compartment c1

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

Name T Helper Cell

SBO:0000290 physical compartment

# 4 Species

This model contains 94 species. The boundary condition of 35 of these species is set to true so that these species' amount cannot be changed by any reaction. Section 10 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
s22	eIFNg	default	$\text{mol} \cdot l^{-1}$		$\Box$
s11	eIL12	default	$\operatorname{mol} \cdot l^{-1}$		
s51	eIL21	default	$\text{mol} \cdot l^{-1}$		
s55	eIL23	default	$\text{mol} \cdot l^{-1}$		$\Box$
s30	eIL4	default	$\text{mol} \cdot l^{-1}$		$\Box$
s87	eTGFb	default	$\mathrm{mol}\cdot\mathrm{l}^{-1}$		
s89	eIL2	default	$\mathrm{mol}\cdot \mathrm{l}^{-1}$		$\Box$
s90	eIL6	default	$\mathrm{mol}\cdot \mathrm{l}^{-1}$		$\Box$
species_1	eIL17	default	$\operatorname{mol} \cdot 1^{-1}$		
species_2	eIL10	default	$\operatorname{mol} \cdot 1^{-1}$		
species_12	eIL18	default	$\operatorname{mol} \cdot 1^{-1}$		
species_13	anti-IL4	default	$\mathrm{mol}\cdot\mathrm{l}^{-1}$	$\square$	
species_14	anti-IFNg	default	$\mathrm{mol}\cdot \mathrm{l}^{-1}$	$\square$	
species_15	pIL4	default	$\mathrm{mol}\cdot \mathrm{l}^{-1}$	$\square$	
species_16	IL18_pool	default	$\mathrm{mol}\cdot\mathrm{l}^{-1}$	$\square$	
species_17	IL12_pool	default	$\operatorname{mol} \cdot \operatorname{l}^{-1}$	$\square$	
species_18	IFNg_pool	default	$\operatorname{mol} \cdot \operatorname{l}^{-1}$	$\square$	
species_19	IL21_pool	default	$\operatorname{mol} \cdot \operatorname{l}^{-1}$	$\square$	
species_20	IL23_pool	default	$\operatorname{mol} \cdot 1^{-1}$	$   \overline{\mathscr{L}} $	
species_21	IL17_pool	default	$\mathrm{mol}\cdot \mathrm{l}^{-1}$	$\square$	
species_22	IL10_pool	default	$\mathrm{mol}\cdot\mathrm{l}^{-1}$	$\square$	

Id

Name

						tion
	species_23	IL6_pool	default	$\text{mol} \cdot l^{-1}$		
	species_24	IL2_pool	default	$\text{mol} \cdot 1^{-1}$	$\overline{\mathbf{Z}}$	
	species_25	TGFb_pool	default	$\text{mol} \cdot 1^{-1}$		
	species_26	IL4_pool	default	$\text{mol} \cdot 1^{-1}$		
	species_27	pIL10	default	$\text{mol} \cdot 1^{-1}$		
	s48	IL6-IL6R	c1	$\text{mol} \cdot 1^{-1}$		
	s47	IL6R	c1	$\text{mol} \cdot 1^{-1}$		
	s46	IL6	c1	$\text{mol} \cdot 1^{-1}$		
Pro	s45	TGFb-TGFbR	c1	$\text{mol} \cdot 1^{-1}$		
Produced by SBML2l <sup>eT</sup> EX	s44	TGFbR	c1	$\text{mol} \cdot 1^{-1}$		
ed	s43	TGFb	c1	$\text{mol} \cdot 1^{-1}$		
by	s38	IL2-IL2R	c1	$\text{mol} \cdot 1^{-1}$		
<u>₩</u>	s37	IL2R	c1	$\text{mol} \cdot 1^{-1}$		
$\leq$	s36	IL2	c1	$\text{mol} \cdot 1^{-1}$		
Ä	s33	IL4-IL4R	c1	$\text{mol} \cdot 1^{-1}$		
$\mathbb{Z}$	s32	IL4R	c1	$\text{mol} \cdot 1^{-1}$		
	s31	IL4	c1	$\text{mol} \cdot 1^{-1}$		$\Box$
	s25	IFNg-IFNgR	c1	$\text{mol} \cdot 1^{-1}$		
	s24	IFNgR	c1	$\text{mol} \cdot 1^{-1}$		
	s14	IL12-IL12R	c1	$\text{mol} \cdot 1^{-1}$		
	s13	IL12R	c1	$\text{mol} \cdot 1^{-1}$		$\Box$
	s12	IL12	c1	$\text{mol} \cdot 1^{-1}$		
	s3	IL18-IL18R	c1	$\text{mol} \cdot 1^{-1}$		$\Box$
	s2	IL18R	c1	$\text{mol} \cdot 1^{-1}$		$\Box$
	s1	IL18	c1	$\text{mol} \cdot 1^{-1}$		
	s52	IL21	c1	$\text{mol} \cdot 1^{-1}$		
	s54	IL21-IL21R	c1	$\text{mol} \cdot l^{-1}$		

Compartment

Constant Boundary Condi-

Derived Unit

Id	Name	Compartment	Derived Unit	Constant	Boundary Condi- tion
s53	IL21R	c1	$\text{mol} \cdot 1^{-1}$	$\Box$	$\Box$
s58	IL23R	c1	$\text{mol} \cdot 1^{-1}$		
s59	IL23-IL23R	c1	$\operatorname{mol} \cdot 1^{-1}$		
s62	IL10R	c1	$\operatorname{mol} \cdot 1^{-1}$		
s63	IL10-IL10R	c1	$\operatorname{mol} \cdot 1^{-1}$		
s65	IRAK1	c1	$\operatorname{mol} \cdot 1^{-1}$		
s10	IRAK1-P	c1	$\operatorname{mol} \cdot 1^{-1}$		
s20	p50/p65 dimer	c1	$\operatorname{mol} \cdot 1^{-1}$		
s67	STAT4	c1	$\operatorname{mol} \cdot 1^{-1}$		
s21	STAT4-P	c1	$\mathrm{mol}\cdot\mathrm{l}^{-1}$		
s68	IFNg	c1	$\mathrm{mol}\cdot\mathrm{l}^{-1}$		
s69	JAK1	c1	$\mathrm{mol}\cdot\mathrm{l}^{-1}$		
s28	JAK1-P	c1	$\operatorname{mol} \cdot 1^{-1}$		
s70	STAT1	c1	$\operatorname{mol} \cdot 1^{-1}$		
s26	STAT1-P	c1	$\operatorname{mol} \cdot 1^{-1}$		
s57	p40/p19 dimer	c1	$\text{mol} \cdot 1^{-1}$		
s73	IL17	c1	$\mathrm{mol}\cdot\mathrm{l}^{-1}$		
s74	IL10	c1	$\mathrm{mol}\cdot\mathrm{l}^{-1}$		
s75	RORgt	c1	$\mathrm{mol}\cdot\mathrm{l}^{-1}$		
<b>s</b> 50	RORgt-ligand	c1	$\text{mol} \cdot 1^{-1}$		
s49	STAT3-P	c1	$\operatorname{mol} \cdot 1^{-1}$		
s76	STAT3	c1	$\operatorname{mol} \cdot 1^{-1}$		$\checkmark$
s39	STAT5-P	c1	$\mathrm{mol}\cdot\mathrm{l}^{-1}$		
s77	STAT5	c1	$\text{mol} \cdot 1^{-1}$		
s78	FOXP3	c1	$\text{mol} \cdot 1^{-1}$		
s79	SOCS1	c1	$\text{mol} \cdot 1^{-1}$		
s29	SOCS1-JAKs	c1	$\text{mol} \cdot 1^{-1}$		

•	Id	Name	Compartment	Derived Unit	Constant	Boundary Condi- tion
	s27	Tbet-P	c1	$\text{mol} \cdot 1^{-1}$		
	s80	Tbet	c1	$\operatorname{mol} \cdot 1^{-1}$		
	s81	GATA3	c1	$\mathrm{mol}\cdot \mathrm{l}^{-1}$		
	s35	GATA3-P	c1	$\mathrm{mol}\cdot \mathrm{l}^{-1}$		
	s34	STAT6-P	c1	$\operatorname{mol} \cdot 1^{-1}$		$\Box$
	s82	STAT6	c1	$\text{mol} \cdot 1^{-1}$		
	s85	PPARg	c1	$\text{mol} \cdot 1^{-1}$		$\exists$
<b>.</b>	s83	L-PPARg	c1	$\text{mol} \cdot 1^{-1}$		$\Box$
Pro	s86	Ligand	c1	$\operatorname{mol} \cdot 1^{-1}$		
duc	s40	acetylated FOXP3	c1	$\mathrm{mol}\cdot \mathrm{l}^{-1}$		
Produced by	species_8	p19	c1	$\operatorname{mol} \cdot 1^{-1}$		
	species_9	p40	c1	$\operatorname{mol} \cdot 1^{-1}$		$\checkmark$
SBMIZIATEX	${ t species\_10}$	p50	c1	$\operatorname{mol} \cdot 1^{-1}$		$\checkmark$
$\leq$	species_11	p65	c1	$\operatorname{mol} \cdot 1^{-1}$		$\checkmark$
Ά	${ t species\_4}$	IL17R	c1	$\operatorname{mol} \cdot 1^{-1}$		
Z.	species_3	IL17-IL17R	c1	$\operatorname{mol} \cdot 1^{-1}$		$\Box$
	species_5	pIFNg	c1	$\mathrm{mol}\cdot \mathrm{l}^{-1}$		
	species_6	pIL21	c1	$\operatorname{mol} \cdot 1^{-1}$		
	${\tt species\_7}$	pIL17	c1	$\text{mol} \cdot 1^{-1}$		

## **5 Parameters**

This model contains five global parameters.

Table 4: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
parameter_1	Hill Coeficient	2.000	
$parameter_2$	BActin	0.009	
$parameter_3$	FOXP3R	0.000	
$parameter_4$	IL17R	0.000	
${\tt ModelValue\_5}$	Initial for BActin	0.009	$\square$

## 6 Initialassignments

This is an overview of twelve initial assignments.

## **6.1 Initialassignment** s22

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

Math [species\_18]

## **6.2 Initialassignment** s11

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

Math [species\_17]

## **6.3 Initialassignment** s51

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

Math [species\_19]

## **6.4 Initialassignment** s55

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

Math [species\_20]

## 6.5 Initialassignment s30

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

Math [species\_26]

## 6.6 Initialassignment s87

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

Math [species\_25]

## 6.7 Initialassignment s89

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

Math [species\_24]

## 6.8 Initialassignment s90

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

Math [species\_23]

## **6.9 Initialassignment** species\_1

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

Math [species\_21]

## **6.10 Initialassignment** species\_2

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

Math [species\_22]

## **6.11 Initialassignment** species\_12

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

Math [species\_16]

## **6.12 Initialassignment ModelValue\_5**

**Derived unit** contains undeclared units

Math parameter\_2

## 7 Function definitions

This is an overview of 16 function definitions.

## 7.1 Function definition function\_1

Name 2 Reactants, 1 Inhibitor

Arguments Vf, r1, r2, K, n, I, Vr, p

**Mathematical Expression** 

$$Vf \cdot r1 \cdot r2 \cdot \frac{K^n}{I^n + K^n} - Vr \cdot p \tag{1}$$

#### 7.2 Function definition function\_2

Name 1 Reactant, 1 Activator

Arguments Vf, r1, A, n, K, Vr, p

**Mathematical Expression** 

$$Vf \cdot r1 \cdot \left(1 + \frac{A^n}{A^n + K^n}\right) - Vr \cdot p$$
 (2)

#### 7.3 Function definition function\_3

Name 2 Reactants, 2 Inhibitors

**Arguments** Vf, r1, r2, K1, n1, I1, K2, n2, I2, Vr, p

**Mathematical Expression** 

$$Vf \cdot r1 \cdot r2 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} - Vr \cdot p \tag{3}$$

#### 7.4 Function definition function\_4

Name 2 Reactants, 2 Inhibitors, 1 Activator

**Arguments** Vf, K1, n1, I1, K2, n2, I2, A, n3, K3, Vr, p, r1, r2

**Mathematical Expression** 

$$Vf \cdot r1 \cdot r2 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \left(1 + \frac{A^{n3}}{A^{n3} + K3^{n3}}\right) - Vr \cdot p \qquad (4)$$

#### 7.5 Function definition function\_6

Name 1 Reactant, 1 Inhibitor, 3 Activators

**Arguments** Vf, r1, K, n, I, A1, n1, K1, A2, n2, K2, A3, n3, K3, Vr, p

#### **Mathematical Expression**

$$Vf \cdot r1 \cdot \frac{K^n}{I^n + K^n} \cdot \left( \frac{A1^{n1}}{A1^{n1} + K1^{n1}} + \frac{A2^{n2}}{A2^{n2} + K2^{n2}} + \frac{A3^{n3}}{A3^{n3} + K3^{n3}} \right) - Vr \cdot p \quad (5)$$

#### **7.6 Function definition** function\_7

Name 1 Reactant, 3 Inhibitors, 3 Activators

**Arguments** Vf, r1, K1, n1, I1, K2, n2, I2, K3, n3, I3, A1, n4, K4, A2, n5, K5, A3, n6, K6, Vr, p

#### **Mathematical Expression**

$$\begin{split} Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \frac{K3^{n3}}{I3^{n3} + K3^{n3}} \\ \cdot \left( \frac{A1^{n4}}{A1^{n4} + K4^{n4}} + \frac{A2^{n5}}{A2^{n5} + K5^{n5}} + \frac{A3^{n6}}{A3^{n6} + K6^{n6}} \right) - Vr \cdot p \end{split} \tag{6}$$

#### 7.7 Function definition function\_8

Name 1 Reactant, 2 Inhibitors, 1 Activator

**Arguments** Vf, r1, K1, n1, I1, K2, n2, I2, A, n3, K3, Vr, p

#### **Mathematical Expression**

$$Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \left(1 + \frac{A^{n3}}{A^{n3} + K3^{n3}}\right) - Vr \cdot p \tag{7}$$

#### 7.8 Function definition function\_9

Name 1 Reactant, 2 Activators

**Arguments** Vf, r1, A1, n1, K1, A2, n2, K2, Vr, p

#### **Mathematical Expression**

$$Vf \cdot r1 \cdot \frac{A1^{n1}}{A1^{n1} + K1^{n1}} \cdot \frac{A2^{n2}}{A2^{n2} + K2^{n2}} - Vr \cdot p \tag{8}$$

#### 7.9 Function definition function\_10

Name 1 Reactant, 3 Inhibitors, 2 Activators

**Arguments** Vf, r1, K1, n1, I1, K2, n2, I2, K3, n3, I3, A1, n4, K4, A2, n5, K5, Vr, p

## **Mathematical Expression**

$$\begin{split} Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \frac{K3^{n3}}{I3^{n3} + K3^{n3}} \\ \cdot \left( \frac{A1^{n4}}{A1^{n4} + K4^{n4}} + \frac{A2^{n5}}{A2^{n5} + K5^{n5}} \right) - Vr \cdot p \end{split} \tag{9}$$

#### 7.10 Function definition function\_12

Name 1 Reactant, 1 Inhibitor, 4 Activators

**Arguments** Vf, r1, K, n, I, A1, n1, K1, A2, n2, K2, A3, n3, K3, A4, n4, K4, Vr, p

#### **Mathematical Expression**

$$\begin{split} Vf \cdot r1 \cdot \frac{K^{n}}{I^{n} + K^{n}} \\ \cdot \left( \frac{A1^{n1}}{A1^{n1} + K1^{n1}} + \frac{A2^{n2}}{A2^{n2} + K2^{n2}} + \frac{A3^{n3}}{A3^{n3} + K3^{n3}} + \frac{A4^{n4}}{A4^{n4} + K4^{n4}} \right) - Vr \cdot p \end{split} \tag{10}$$

#### 7.11 Function definition function\_14

Name 2 Reactants, 1 Inhibitor, 1 Activator

**Arguments** Vf, r1, r2, K, n, I, A, n1, K1, Vr, p

#### **Mathematical Expression**

$$Vf \cdot r1 \cdot r2 \cdot \frac{K^n}{I^n + K^n} \cdot \left(1 + \frac{A^{n1}}{A^{n1} + K1^{n1}}\right) - Vr \cdot p \tag{11}$$

## 7.12 Function definition function\_11

Name 1 Reactant, 3 Inhibitors, 4 Activators

**Arguments** Vf, r1, K1, n1, I1, K2, n2, I2, K3, n3, I3, A1, n4, K4, A2, n5, K5, A3, n6, K6, A4, n7, K7, Vr, p

#### **Mathematical Expression**

$$Vf \cdot r1 \cdot \frac{K1^{n1}}{11^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{12^{n2} + K2^{n2}} \cdot \frac{K3^{n3}}{13^{n3} + K3^{n3}}$$

$$\cdot \left( \frac{A1^{n4}}{A1^{n4} + K4^{n4}} + \frac{A2^{n5}}{A2^{n5} + K5^{n5}} + \frac{A3^{n6}}{A3^{n6} + K6^{n6}} + \frac{A4^{n7}}{A4^{n7} + K7^{n7}} \right) - Vr \cdot p$$

$$(12)$$

## 7.13 Function definition function\_5

Name 1 Reactant, 2 Inhibitors, 2 Activators

**Arguments** Vf, r1, K1, n1, I1, K2, n2, I2, A1, n3, K3, A2, n4, K4, Vr, p

#### **Mathematical Expression**

$$Vf \cdot r1 \cdot \frac{K1^{n1}}{11^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{12^{n2} + K2^{n2}} \cdot \left(\frac{A1^{n3}}{A1^{n3} + K3^{n3}} + \frac{A2^{n4}}{A2^{n4} + K4^{n4}}\right) - Vr \cdot p \quad (13)$$

#### 7.14 Function definition function\_13

Name 1 Reactant, 2 Inhibitors, 3 Activators

**Arguments** Vf, r1, K1, n1, I1, K2, n2, I2, A1, n3, K3, A2, n4, K4, A3, n5, K5, Vr, p

### **Mathematical Expression**

$$\begin{split} Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \\ \cdot \left( \frac{A1^{n3}}{A1^{n3} + K3^{n3}} + \frac{A2^{n4}}{A2^{n4} + K4^{n4}} + \frac{A3^{n5}}{A3^{n5} + K5^{n5}} \right) - Vr \cdot p \end{split} \tag{14}$$

## 7.15 Function definition function\_15

Name 1 Reactant, 1 inhibitor, 1 activator

**Arguments** Vf, r1, K, n, I, A1, n1, K1, Vr, p

#### **Mathematical Expression**

$$Vf \cdot r1 \cdot \frac{K^{n}}{I^{n} + K^{n}} \cdot \frac{A1^{n1}}{A1^{n1} + K1^{n1}} - Vr \cdot p \tag{15}$$

#### 7.16 Function definition function\_16

Name Pool coupling

Arguments V, pool, n, ext, k

**Mathematical Expression** 

$$V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right)$$
 (16)

## 8 Rules

This is an overview of two rules.

## 8.1 Rule parameter\_3

Rule parameter\_3 is an assignment rule for parameter parameter\_3:

$$parameter\_3 = \frac{[s40]}{ModelValue\_5}$$
 (17)

## 8.2 Rule parameter\_4

Rule parameter\_4 is an assignment rule for parameter parameter\_4:

$$parameter\_4 = \frac{[s73]}{ModelValue\_5}$$
 (18)

# 9 Reactions

This model contains 53 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

			Tuble 5. 6 verview of all reactions	
N⁰	Id	Name	Reaction Equation	SBO
1	re2	re2	species_12+s2 $\stackrel{\text{s34, species}\_12, s2, s34, s3}{\longleftarrow}$ s3	
2	re3	re3	$s65 \rightleftharpoons s3, s65, s3, s10$	
3	re6	re6	$s12 \xrightarrow{s12} s11$	
4	re8	re8	$s11 + s13 \xrightarrow{s34, s83, s11, s13, s34, s83, s14} s14$	
5	re9	re9	species_10+species_11	
6	re10	re10	s67 <u>\$54, \$35, \$14, \$59, \$67, \$54, \$35, \$14, \$59,</u>	$\stackrel{\underline{s21}}{\longrightarrow} s21$
7	re11	re11	$s22 + s24 = \frac{s29, s22, s24, s29, s25}{s25}$	
8	re12	re12	$s68 \xrightarrow{s68} s22$	
9	re13	re13	s70 \(\frac{\s83, \s54, \s25, \s28, \s70, \s83, \s54, \s25, \s28, \rangle}{\sigma}	$\stackrel{s26}{=}$ s26
10	re14	re14	species_5 species_14, s49, s83, s21, s20, s27, s	
11	re15	re15	s80 <u>s45, s49, s21, s26, s27, s80, s45, s49, s21,</u>	<u>\$26, \$27</u> <u>\$27</u>
12	re16	re16	$s69 = \frac{s83, s29, s25, s69, s83, s29, s25, s28}{5} s28$	
13	re17	re17	$s79 = \frac{s26, s27, s79, s26, s27, s29}{s29}$	
14	re18	re18	$s31 \xrightarrow{s31} s30$	
15	re19	re19	$s30 + s32 \stackrel{\underline{s29, s30, s32, s29, s33}}{\longleftarrow} s33$	

Nº	Id	Name	Reaction Equation	SBO
16	re20	re20	species_15 = \$26, species_13, s31, s35, s39, species_15	, s26, species_13, s31, s35, s39
17	re23	re23	s81 <u>\$27, \$49, \$45, \$34, \$83, \$81, \$27, \$49, \$45, \$34, \$</u>	83, s35 s35
18	re24	re24	s82 <u>s25, s33, s82, s25, s33, s34</u> s34	
19	re25	re25	$889 + 837 \xrightarrow{889, 837, 838} 838$	
20	re26	re26	$877 = \frac{838, 877, 838, 839}{2000} 839$	
21	re27	re27	s78 = s34, s49, s48, s45, s26, s39, s83, s78, s34, s49, s	48, s45, s26, s39, s83, s40 s40
22	re28	re28	$887 + 844 \xrightarrow{887, 844, 845} 845$	
23	re29	re29	$s90 + s47 = \frac{s90, s47, s48}{s48} s48$	10 10
24	re30	re30	s76 \$83, s54, s59, s63, s48, s76, s83, s54, s59, s63, s	48, s49 ——→ s49
25	re31	re31	\$75 \(\frac{\section{40, \$83, \$49, \$45, \$75, \$40, \$83, \$49, \$45, \$50}{30, \$51, \$52, \$20, \$20, \$10, \$50, \$20, \$10, \$10, \$10, \$10, \$10, \$10, \$10, \$1	50
26	re32	re32	s51+s53 = s39, species_3, s51, s53, s39, species_3, s5	4 ⇒ s54
27	re33	re33	$s52 = \frac{s52}{55} s51$	
28	re34	re34	$s55 \xrightarrow{s55, s57} s57$	
29	re35	re35	$s57 + s58 \xrightarrow{s27, s57, s58, s27, s59} s59$	02 40 57
30	re36	re36	species_9+species_8	$\xrightarrow{883, 848, 857} 857$
31	re37	re37	species_6 $\stackrel{\text{s49, species}\_6, \text{s49, s52}}{\rightleftharpoons}$ s52	
32	re39	re38	species_2 + s62 $\rightleftharpoons$ species_2, s62, s63	
33	re42	re42	$885 + 886 = \frac{825, 833, 885, 886, 825, 833, 883}{200} $	
34	re44	re44	$s43 \xrightarrow{s43} s87$	

No	Id	Name	Reaction Equation SBO
35	re45	re45	$s36 \xrightarrow{s36} s89$
36	re46	re46	$s46 \xrightarrow{s46} s90$
37	${\tt reaction\_1}$	re48	$s74 \xrightarrow{s74} species_2$
38	${\tt reaction\_2}$	re49	$s73 \xrightarrow{s73} species_1$
39	${\tt reaction\_3}$	re50	species_1 + species_4 species_4, species_3 species_3
40	${\tt reaction\_4}$	re47	$s1 \xrightarrow{S1} species 12$
41	reaction_5	re51	species_7 $\frac{$83, $59, $50, $21, $pecies_7, $83, $59, $50, $21, $73}{$}$ \$73
42	reaction_6	IL18 pool	species_16 species_12 species_12
43	reaction_7	IL12 pool	species_17 $\stackrel{\text{species}\_17, \text{ s}11}{\longleftarrow}$ s11
44	reaction_8	IFNg pool	species_ $18 \xrightarrow{\text{species}\_18, s22} s22$
45	reaction_9	IL21 pool	species_19 $\stackrel{\text{species}\_19, s51}{\longleftarrow}$ s51
46	reaction_10	IL23 pool	species_20 $\rightleftharpoons$ species_5 s55
47	reaction_11	IL17 pool	species_21 $\stackrel{\text{species}\_21, s73}{\longleftarrow}$ s73
48	reaction_12	IL10 pool	species_22 species_2 species_2
49	reaction_13	IL6 pool	species_23 species_23, s90 s90
50	reaction_14	IL2 pool	species_24 species_24, s89 s89
51	reaction_15	TGFb pool	species_25 $\rightleftharpoons$ species_25, s87 $\rightleftharpoons$ s87
52	reaction_16	IL4 pool	species_26, $\stackrel{\text{species}\_26, s30}{=}$ s30
	reaction_17	re52	species_27

#### 9.1 Reaction re2

This is a reversible reaction of two reactants forming one product influenced by five modifiers.

Name re2

## **Reaction equation**

species\_12 + s2 
$$\stackrel{\text{s34, species}_12, s2, s34, s3}{\longleftarrow}$$
 s3 (19)

#### **Reactants**

Table 6: Properties of each reactant.

Id	Name	SBO
species_12 s2	eIL18 IL18R	

#### **Modifiers**

Table 7: Properties of each modifier.

Id	Name	SBO
s34	STAT6-P	
species_12	eIL18	
s2	IL18R	
s34	STAT6-P	
s3	IL18-IL18R	

#### **Product**

Table 8: Properties of each product.

Id	Name	SBO
s3	IL18-IL18R	

## **Kinetic Law**

$$v_1 = \text{function\_1}(Vf, [\text{species\_12}], [\text{s2}], K, \text{parameter\_1}, [\text{s34}], Vr, [\text{s3}])$$
 (20)

$$function\_1\left(Vf,r1,r2,K,n,I,Vr,p\right) = Vf\cdot r1\cdot r2\cdot \frac{K^n}{I^n+K^n} - Vr\cdot p \tag{21}$$

Table 9: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	$\overline{Z}$
K	K	0.138	$\square$
Vr	Vr	0.100	$\square$

#### 9.2 Reaction re3

This is a reversible reaction of one reactant forming one product influenced by four modifiers.

Name re3

## **Reaction equation**

$$s65 \stackrel{\underline{s3, s65, s3, s10}}{\longleftarrow} s10$$
 (22)

#### Reactant

Table 10: Properties of each reactant.

Id	Name	SBO
s65	IRAK1	

#### **Modifiers**

Table 11: Properties of each modifier.

Id	Name	SBO
s3	IL18-IL18R	
s65	IRAK1	
s3	IL18-IL18R	
s10	IRAK1-P	

## **Product**

Table 12: Properties of each product.

Id	Name	SBO
s10	IRAK1-P	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_2 = \text{vol}(c1) \cdot \text{function}_2(Vf, [s65], [s3], \text{parameter}_1, K, Vr, [s10])$$
 (23)

$$function_{-}2\left(Vf,r1,A,n,K,Vr,p\right) = Vf \cdot r1 \cdot \left(1 + \frac{A^n}{A^n + K^n}\right) - Vr \cdot p \tag{24}$$

$$function\_2\left(Vf,r1,A,n,K,Vr,p\right) = Vf \cdot r1 \cdot \left(1 + \frac{A^n}{A^n + K^n}\right) - Vr \cdot p \tag{25}$$

Table 13: Properties of each parameter.

		F	
Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	
K	K	2.017	$\square$
Vr	Vr	0.100	$\square$

#### 9.3 Reaction re6

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

Name re6

#### **Reaction equation**

$$s12 \xrightarrow{s12} s11 \tag{26}$$

## Reactant

Table 14: Properties of each reactant.

Id	Name	SBO
s12	IL12	

## **Modifier**

Table 15: Properties of each modifier.

Id	Name	SBO
s12	IL12	

## **Product**

Table 16: Properties of each product.

Id	Name	SBO
s11	eIL12	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_3 = \mathbf{k}1 \cdot [\mathbf{s}12] \tag{27}$$

Table 17: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k1	k1	0.1	

## 9.4 Reaction re8

This is a reversible reaction of two reactants forming one product influenced by seven modifiers.

Name re8

## **Reaction equation**

$$s11 + s13 = \frac{s34, s83, s11, s13, s34, s83, s14}{s14} s14$$
 (28)

#### **Reactants**

Table 18: Properties of each reactant.

Id	Name	SBO
s11	eIL12	

Id	Name	SBO
s13	IL12R	

#### **Modifiers**

Table 19: Properties of each modifier.

Id	Name	SBO
s34	STAT6-P	
s83	L-PPARg	
s11	eIL12	
s13	IL12R	
s34	STAT6-P	
s83	L-PPARg	
s14	IL12-IL12R	

#### **Product**

Table 20: Properties of each product.

Id	Name	SBO
s14	IL12-IL12R	

#### **Kinetic Law**

$$v_4 = \text{function}_3(Vf, [s11], [s13], K1, \text{parameter}_1, [s34], K2, \text{parameter}_1, [s83], Vr, [s14])$$
 (29)

$$\text{function\_3} \left( Vf, r1, r2, K1, n1, I1, K2, n2, I2, Vr, p \right) = Vf \cdot r1 \cdot r2 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} - Vr \cdot p$$
 (30)

Table 21: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	
K1	K1	2.946	
K2	K2	0.744	
Vr	Vr	0.100	$\square$

## 9.5 Reaction re9

This is a reversible reaction of two reactants forming one product influenced by nine modifiers.

#### Name re9

## **Reaction equation**

$$species\_10 + species\_11 \xrightarrow{s83, s40, s10, s83, s40, s10, s20, species\_10, species\_11} s20 \tag{31}$$

#### **Reactants**

Table 22: Properties of each reactant.

Id	Name	SBO
species_10 species_11	p50 p65	

#### **Modifiers**

Table 23: Properties of each modifier.

	1	
Id	Name	SBO
s83	L-PPARg	
s40	acetylated FOXP3	
s10	IRAK1-P	
s83	L-PPARg	
s40	acetylated FOXP3	
s10	IRAK1-P	
s20	p50/p65 dimer	
species_10	p50	
species_11	p65	

#### **Product**

Table 24: Properties of each product.

Id	Name	SBO
s20	p50/p65 dimer	

#### **Kinetic Law**

#### **Derived unit** contains undeclared units

$$v_5 = \text{vol}(c1) \cdot \text{function}\_4(Vf, K1, \text{parameter}\_1, [s83], K2, \text{parameter}\_1, [s40], [s10], parameter}\_1, K3, Vr, [s20], [species}\_10], [species}\_11])$$
(32)

$$\begin{split} &\text{function\_4}\left(Vf, K1, n1, I1, K2, n2, I2, A, n3, K3, Vr, p, r1, r2\right) \\ &= Vf \cdot r1 \cdot r2 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \left(1 + \frac{A^{n3}}{A^{n3} + K3^{n3}}\right) - Vr \cdot p \end{split} \tag{33}$$

$$\begin{split} &\text{function\_4} \left( Vf, K1, n1, I1, K2, n2, I2, A, n3, K3, Vr, p, r1, r2 \right) \\ &= Vf \cdot r1 \cdot r2 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \left( 1 + \frac{A^{n3}}{A^{n3} + K3^{n3}} \right) - Vr \cdot p \end{split} \tag{34}$$

Table 25: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	
K1	K1	0.100	
K2	K2	98.948	
КЗ	K3	0.054	
Vr	Vr	0.100	$\checkmark$

#### 9.6 Reaction re10

This is a reversible reaction of one reactant forming one product influenced by ten modifiers.

#### Name re10

#### **Reaction equation**

$$s67 \stackrel{s54, s35, s14, s59, s67, s54, s35, s14, s59, s21}{\longleftarrow} s21$$
 (35)

#### Reactant

Table 26: Properties of each reactant.

Id	Name	SBO
s67	STAT4	

#### **Modifiers**

Table 27: Properties of each modifier.

Id	Name	SBO
s54	IL21-IL21R	
<b>s</b> 35	GATA3-P	
s14	IL12-IL12R	
s59	IL23-IL23R	
s67	STAT4	
s54	IL21-IL21R	
s35	GATA3-P	
s14	IL12-IL12R	
s59	IL23-IL23R	
s21	STAT4-P	

#### **Product**

Table 28: Properties of each product.

Id	Name	SBO
s21	STAT4-P	

#### **Kinetic Law**

$$v_6 = vol\left(c1\right) \cdot function\_5\left(Vf, [s67], K1, parameter\_1, [s54], K2, parameter\_1, [s35], [s14], \\ parameter\_1, K3, [s59], parameter\_1, K4, Vr, [s21] \right)$$

$$\begin{split} &\text{function\_5} \left( Vf, r1, K1, n1, I1, K2, n2, I2, A1, n3, K3, A2, n4, K4, Vr, p \right) \\ &= Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \left( \frac{A1^{n3}}{A1^{n3} + K3^{n3}} + \frac{A2^{n4}}{A2^{n4} + K4^{n4}} \right) - Vr \cdot p \end{split} \tag{37}$$

$$\begin{split} &\text{function\_5} \left( Vf, r1, K1, n1, I1, K2, n2, I2, A1, n3, K3, A2, n4, K4, Vr, p \right) \\ &= Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \left( \frac{A1^{n3}}{A1^{n3} + K3^{n3}} + \frac{A2^{n4}}{A2^{n4} + K4^{n4}} \right) - Vr \cdot p \end{split} \tag{38}$$

Table 29: Properties of each parameter.

		1 1	
Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	$ \mathcal{A} $
K1	K1	0.125	
K2	K2	0.897	
КЗ	K3	0.031	
K4	K4	66.617	$ \overline{\checkmark} $
Vr	Vr	0.100	$\overline{\mathbf{Z}}$

## 9.7 Reaction re11

This is a reversible reaction of two reactants forming one product influenced by five modifiers.

#### Name rel1

## **Reaction equation**

$$s22 + s24 \xrightarrow{s29, s22, s24, s29, s25} s25$$
 (39)

## **Reactants**

Table 30: Properties of each reactant.

Name	SBO
eIFNg IFNgR	
	- 1 (01110

#### **Modifiers**

Table 31: Properties of each modifier.

Id	Name	SBO
s29	SOCS1-JAKs	
s22	eIFNg	
s24	IFNgR	
s29	SOCS1-JAKs	
s25	IFNg-IFNgR	

## **Product**

Table 32: Properties of each product.

Id	Name	SBO
s25	IFNg-IFNgR	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_7 = \text{function}_1(Vf, [s22], [s24], K, \text{parameter}_1, [s29], Vr, [s25])$$
 (40)

$$function\_1\left(Vf,r1,r2,K,n,I,Vr,p\right) = Vf\cdot r1\cdot r2\cdot \frac{K^n}{I^n+K^n} - Vr\cdot p \tag{41} \label{eq:41}$$

Table 33: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	
K	K	0.264	$ \overline{\checkmark} $
Vr	Vr	0.100	$\checkmark$

#### 9.8 Reaction re12

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

## Name re12

## **Reaction equation**

$$s68 \xrightarrow{s68} s22 \tag{42}$$

#### Reactant

Table 34: Properties of each reactant.

Id	Name	SBO
s68	IFNg	

#### **Modifier**

Table 35: Properties of each modifier.

Id	Name	SBO
s68	IFNg	

#### **Product**

Table 36: Properties of each product.

Id	Name	SBO
s22	eIFNg	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_8 = k1 \cdot [s68] \tag{43}$$

Table 37: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k1	k1	0.1	$\overline{Z}$

#### 9.9 Reaction re13

This is a reversible reaction of one reactant forming one product influenced by ten modifiers.

#### Name re13

## **Reaction equation**

$$s70 = \frac{s83, s54, s25, s28, s70, s83, s54, s25, s28, s26}{s26}$$
 s26 (44)

#### Reactant

Table 38: Properties of each reactant.

Id	Name	SBO
s70	STAT1	

#### **Modifiers**

Table 39: Properties of each modifier.

Id	Name	SBO
s83	L-PPARg	
s54	IL21-IL21R	
s25	IFNg-IFNgR	
s28	JAK1-P	
s70	STAT1	
s83	L-PPARg	
s54	IL21-IL21R	
s25	IFNg-IFNgR	
s28	JAK1-P	
s26	STAT1-P	

#### **Product**

Table 40: Properties of each product.

Id	Name	SBO
s26	STAT1-P	

#### **Kinetic Law**

$$v_9 = vol\left(c1\right) \cdot function\_6\left(Vf, [s70], K, parameter\_1, [s83], [s54], parameter\_1, K1, [s25], \\ parameter\_1, K2, [s28], parameter\_1, K3, Vr, [s26] \right)$$

$$\begin{split} &\text{function\_6}\left(Vf, r1, K, n, I, A1, n1, K1, A2, n2, K2, A3, n3, K3, Vr, p\right) \\ &= Vf \cdot r1 \cdot \frac{K^n}{I^n + K^n} \cdot \left(\frac{A1^{n1}}{A1^{n1} + K1^{n1}} + \frac{A2^{n2}}{A2^{n2} + K2^{n2}} + \frac{A3^{n3}}{A3^{n3} + K3^{n3}}\right) - Vr \cdot p \end{split} \tag{46}$$

$$\begin{split} &\text{function\_6}\left(Vf,r1,K,n,I,A1,n1,K1,A2,n2,K2,A3,n3,K3,Vr,p\right) \\ &= Vf \cdot r1 \cdot \frac{K^n}{I^n + K^n} \cdot \left(\frac{A1^{n1}}{A1^{n1} + K1^{n1}} + \frac{A2^{n2}}{A2^{n2} + K2^{n2}} + \frac{A3^{n3}}{A3^{n3} + K3^{n3}}\right) - Vr \cdot p \end{split} \tag{47}$$

Table 41: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	lacksquare
K	K	0.100	$\overline{Z}$
K1	K1	5.044	$   \overline{\mathscr{A}} $
K2	K2	0.071	$   \overline{\mathscr{A}} $
КЗ	K3	14.978	
Vr	Vr	0.100	$   \overline{\mathscr{A}} $

## 9.10 Reaction re14

This is a reversible reaction of one reactant forming one product influenced by 14 modifiers.

#### Name re14

## **Reaction equation**

#### Reactant

Table 42: Properties of each reactant.

Id	Name	SBO
species_5	pIFNg	

## **Modifiers**

Table 43: Properties of each modifier.

Id	Name	SBO
species_14	anti-IFNg	
s49	STAT3-P	
s83	L-PPARg	
s21	STAT4-P	
s20	p50/p65 dimer	
s27	Tbet-P	
species_5	pIFNg	
${\tt species\_14}$	anti-IFNg	
s49	STAT3-P	

Id	Name	SBO
s83	L-PPARg	
s21	STAT4-P	
s20	p50/p65 dimer	
s27	Tbet-P	
s68	IFNg	

#### **Product**

Table 44: Properties of each product.

Id	Name	SBO
s68	IFNg	

#### **Kinetic Law**

$$\begin{aligned} v_{10} &= vol\left(c1\right) \cdot function\_7\left(Vf, [species\_5], K1, parameter\_1, [species\_14], K2, parameter\_1, \\ & [s49], K3, parameter\_1, [s83], [s21], parameter\_1, K4, [s20], parameter\_1, K5, [s27], \\ & parameter\_1, K6, Vr, [s68]\right) \end{aligned} \tag{49}$$

$$\begin{split} &\text{function\_7} \left( Vf, r1, K1, n1, I1, K2, n2, I2, K3, n3, I3, A1, n4, K4, A2, n5, K5, A3, n6, K6, Vr, p \right) \\ &= Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \frac{K3^{n3}}{I3^{n3} + K3^{n3}} \end{split}$$

$$= Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \frac{K3^{n3}}{I3^{n3} + K3^{n3}} \cdot \left(\frac{A1^{n4}}{A1^{n4} + K4^{n4}} + \frac{A2^{n5}}{A2^{n5} + K5^{n5}} + \frac{A3^{n6}}{A3^{n6} + K6^{n6}}\right) - Vr \cdot p$$
(50)

$$\begin{split} & \text{function\_7} \left( Vf, r1, K1, n1, I1, K2, n2, I2, K3, n3, I3, A1, n4, K4, A2, n5, K5, A3, n6, K6, Vr, p \right) \\ &= Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \frac{K3^{n3}}{I3^{n3} + K3^{n3}} \\ & \cdot \left( \frac{A1^{n4}}{A1^{n4} + K4^{n4}} + \frac{A2^{n5}}{A2^{n5} + K5^{n5}} + \frac{A3^{n6}}{A3^{n6} + K6^{n6}} \right) - Vr \cdot p \end{split}$$

Table 45: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	lacksquare
K1	K1	0.502	$   \overline{\mathbf{A}} $
K2	K2	0.812	$   \overline{\mathbf{A}} $
КЗ	K3	0.100	
K4	K4	0.001	
K5	K5	100.000	$   \overline{\mathbf{A}} $
K6	K6	0.231	$   \overline{\mathbf{A}} $
Vr	Vr	0.100	$\checkmark$

## 9.11 Reaction re15

This is a reversible reaction of one reactant forming one product influenced by eleven modifiers.

#### Name re15

## **Reaction equation**

$$s80 = \frac{s45, s49, s21, s26, s27, s80, s45, s49, s21, s26, s27}{s27} s27$$
 (52)

#### Reactant

Table 46: Properties of each reactant.

Id	Name	SBO
s80	Tbet	

#### **Modifiers**

Table 47: Properties of each modifier.

Id	Name	SBO
s45	TGFb-TGFbR	
s49	STAT3-P	
s21	STAT4-P	
s26	STAT1-P	
s27	Tbet-P	
s80	Tbet	
s45	TGFb-TGFbR	
s49	STAT3-P	

Id	Name	SBO
s21	STAT4-P	
s26	STAT1-P	
s27	Tbet-P	

#### **Product**

Table 48: Properties of each product.

Id	Name	SBO
s27	Tbet-P	

#### Kinetic Law

$$\begin{aligned} \nu_{11} &= vol\left(c1\right) \cdot function\_13\left(Vf, [s80], K1, parameter\_1, [s45], K2, parameter\_1, [s49], \\ & [s21], parameter\_1, K3, [s26], parameter\_1, K4, [s27], parameter\_1, K5, Vr, \\ & [s27]) \end{aligned} \tag{53}$$

$$\begin{split} & \text{function\_13} \left( Vf, r1, K1, n1, I1, K2, n2, I2, A1, n3, K3, A2, n4, K4, A3, n5, K5, Vr, p \right) \\ & = Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \left( \frac{A1^{n3}}{A1^{n3} + K3^{n3}} + \frac{A2^{n4}}{A2^{n4} + K4^{n4}} + \frac{A3^{n5}}{A3^{n5} + K5^{n5}} \right) - Vr \cdot p \end{split} \tag{54}$$

$$\begin{split} &\text{function\_13} \left( Vf, r1, K1, n1, I1, K2, n2, I2, A1, n3, K3, A2, n4, K4, A3, n5, K5, Vr, p \right) \\ &= Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \left( \frac{A1^{n3}}{A1^{n3} + K3^{n3}} + \frac{A2^{n4}}{A2^{n4} + K4^{n4}} + \frac{A3^{n5}}{A3^{n5} + K5^{n5}} \right) - Vr \cdot p \end{split} \tag{55}$$

Table 49: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	$\square$
K1	K1	0.917	
K2	K2	1.335	
КЗ	K3	3.588	$\square$
K4	K4	0.728	$\square$
K5	K5	6.978	$\square$
Vr	Vr	0.100	

## 9.12 Reaction re16

This is a reversible reaction of one reactant forming one product influenced by eight modifiers.

Name re16

## **Reaction equation**

$$s69 = \frac{s83, s29, s25, s69, s83, s29, s25, s28}{569} s28$$
 (56)

#### Reactant

Table 50: Properties of each reactant.

Id	Name	SBO
s69	JAK1	

## **Modifiers**

Table 51: Properties of each modifier.

Id	Name	SBO
s83	L-PPARg	
s29	SOCS1-JAKs	
s25	IFNg-IFNgR	
s69	JAK1	
s83	L-PPARg	
s29	SOCS1-JAKs	
s25	IFNg-IFNgR	
s28	JAK1-P	

#### **Product**

Table 52: Properties of each product.

Id	Name	SBO
s28	JAK1-P	

## **Kinetic Law**

$$v_{12} = vol(c1) \cdot function\_8 (Vf, [s69], K1, parameter\_1, [s83], K2, parameter\_1, [s29], [s25], \\ parameter\_1, K3, Vr, [s28])$$
 (57)

$$\begin{split} &\text{function\_8}\left(Vf, r1, K1, n1, I1, K2, n2, I2, A, n3, K3, Vr, p\right) \\ &= Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \left(1 + \frac{A^{n3}}{A^{n3} + K3^{n3}}\right) - Vr \cdot p \end{split} \tag{58}$$

$$\begin{split} &\text{function\_8}\left(Vf, r1, K1, n1, I1, K2, n2, I2, A, n3, K3, Vr, p\right) \\ &= Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \left(1 + \frac{A^{n3}}{A^{n3} + K3^{n3}}\right) - Vr \cdot p \end{split} \tag{59}$$

Table 53: Properties of each parameter.

Id         Name         SBO         Value         Unit         Constant           Vf         Vf         0.100         ✓           K1         K1         0.100         ✓           K2         K2         0.004         ✓           K3         K3         99.987         ✓           Vr         Vr         0.100         ✓			1 1	
K1       K1       0.100       ✓         K2       K2       0.004       ✓         K3       K3       99.987       ✓	Id	Name	SBO Value Unit	Constant
K2       0.004         K3       K3         99.987       ✓	Vf	Vf	0.100	$\overline{Z}$
K3 K3 99.987 ✓	K1	K1	0.100	
	K2	K2	0.004	
Vr	КЗ	K3	99.987	
	Vr	Vr	0.100	

#### 9.13 Reaction re17

This is a reversible reaction of one reactant forming one product influenced by six modifiers.

#### Name re17

#### **Reaction equation**

$$s79 = \frac{s26, s27, s79, s26, s27, s29}{s29} s29$$
 (60)

#### Reactant

Table 54: Properties of each reactant.

Id	Name	SBO
s79	SOCS1	

#### **Modifiers**

Table 55: Properties of each modifier.

Id	Name	SBO
s26	STAT1-P	
s27	Tbet-P	
s79	SOCS1	
s26	STAT1-P	
s27	Tbet-P	
s29	SOCS1-JAKs	

#### **Product**

Table 56: Properties of each product.

Id	Name	SBO
s29	SOCS1-JAKs	

#### **Kinetic Law**

$$v_{13} = vol(c1) \cdot function\_9(Vf, [s79], [s26], parameter\_1, K1, [s27], parameter\_1, K2, Vr, [s29])$$
(61)

$$function\_9 \left( Vf, r1, A1, n1, K1, A2, n2, K2, Vr, p \right) = Vf \cdot r1 \cdot \frac{A1^{n1}}{A1^{n1} + K1^{n1}} \cdot \frac{A2^{n2}}{A2^{n2} + K2^{n2}} - Vr \cdot p$$
 (62)

$$function\_9 \left( Vf, r1, A1, n1, K1, A2, n2, K2, Vr, p \right) = Vf \cdot r1 \cdot \frac{A1^{n1}}{A1^{n1} + K1^{n1}} \cdot \frac{A2^{n2}}{A2^{n2} + K2^{n2}} - Vr \cdot p \right)$$
 (63)

Table 57: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	$\overline{\square}$
K1	K1	7.838	$\overline{\mathbf{Z}}$
K2	K2	0.667	$   \overline{\mathbf{Z}} $
Vr	Vr	0.100	$\square$

# 9.14 Reaction re18

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

Name re18

# **Reaction equation**

$$s31 \xrightarrow{s31} s30 \tag{64}$$

#### Reactant

Table 58: Properties of each reactant.

Id	Name	SBO
s31	IL4	

#### **Modifier**

Table 59: Properties of each modifier.

Id	Name	SBO
s31	IL4	

#### **Product**

Table 60: Properties of each product.

Id	Name	SBO
s30	eIL4	

## **Kinetic Law**

$$v_{14} = k1 \cdot [s31] \tag{65}$$

Table 61: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k1	<b>k</b> 1	0.1	

# 9.15 Reaction re19

This is a reversible reaction of two reactants forming one product influenced by five modifiers.

Name re19

# **Reaction equation**

$$s30 + s32 \xrightarrow{s29, s30, s32, s29, s33} s33$$
 (66)

## **Reactants**

Table 62: Properties of each reactant.

Id	Name	SBO
s30	eIL4	
s32	IL4R	

## **Modifiers**

Table 63: Properties of each modifier.

Id	Name	SBO
s29	SOCS1-JAKs	
<b>s</b> 30	eIL4	
s32	IL4R	
s29	SOCS1-JAKs	
s33	IL4-IL4R	

## **Product**

Table 64: Properties of each product.

Id	Name	SBO
s33	IL4-IL4R	

## **Kinetic Law**

$$v_{15} = \text{function}_{-1}(Vf, [s30], [s32], K, \text{parameter}_{-1}, [s29], Vr, [s33])$$
 (67)

$$function\_1\left(Vf,r1,r2,K,n,I,Vr,p\right) = Vf\cdot r1\cdot r2\cdot \frac{K^n}{I^n+K^n} - Vr\cdot p \tag{68} \label{eq:68}$$

Table 65: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	$ \overline{\mathbf{Z}} $
K	K	13.066	
Vr	Vr	0.100	

## 9.16 Reaction re20

This is a reversible reaction of one reactant forming one product influenced by eleven modifiers.

Name re20

## **Reaction equation**

#### Reactant

Table 66: Properties of each reactant.

Id	Name	SBO
species_15	pIL4	

## **Modifiers**

Table 67: Properties of each modifier.

Id	Name	SBO
	TVallic	эво
s26	STAT1-P	
species_13	anti-IL4	
s31	IL4	
s35	GATA3-P	
s39	STAT5-P	
species_15	pIL4	
s26	STAT1-P	
species_13	anti-IL4	
s31	IL4	

Id	Name	SBO
s35	GATA3-P	
s39	STAT5-P	

#### **Product**

Table 68: Properties of each product.

Id	Name	SBO
s31	IL4	

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{16} = \text{function\_13} \text{ (Vf, [species\_15], K1, parameter\_1, [s26], K2, parameter\_1, [species\_13], } [s31], parameter\_1, K3, [s35], parameter\_1, K4, [s39], parameter\_1, K5, Vr, [s31])$$
 (70)

$$\begin{split} & \text{function\_13} \left( Vf, r1, K1, n1, I1, K2, n2, I2, A1, n3, K3, A2, n4, K4, A3, n5, K5, Vr, p \right) \\ &= Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \left( \frac{A1^{n3}}{A1^{n3} + K3^{n3}} + \frac{A2^{n4}}{A2^{n4} + K4^{n4}} + \frac{A3^{n5}}{A3^{n5} + K5^{n5}} \right) - Vr \cdot p \end{split} \tag{71}$$

Table 69: Properties of each parameter.

		1	
Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	$\overline{Z}$
K1	K1	0.210	
K2	K2	56.345	
КЗ	K3	98.037	
K4	K4	0.856	
K5	K5	4.327	
Vr	Vr	0.100	$ \overline{\checkmark} $

#### 9.17 Reaction re23

This is a reversible reaction of one reactant forming one product influenced by twelve modifiers.

# Name re23

# **Reaction equation**

$$s81 \stackrel{\underline{s27, s49, s45, s34, s83, s81, s27, s49, s45, s34, s83, s35}}{} s35$$
 (72)

## Reactant

Table 70: Properties of each reactant.

Id	Name	SBO
s81	GATA3	

# **Modifiers**

Table 71: Properties of each modifier.

Id	Name	SBO
s27	Tbet-P	
s49	STAT3-P	
s45	TGFb-TGFbR	
s34	STAT6-P	
s83	L-PPARg	
s81	GATA3	
s27	Tbet-P	
s49	STAT3-P	
s45	TGFb-TGFbR	
s34	STAT6-P	
s83	L-PPARg	
s35	GATA3-P	

# **Product**

Table 72: Properties of each product.

Id	Name	SBO
s35	GATA3-P	

#### **Kinetic Law**

$$v_{17} = \text{vol}(c1) \cdot \text{function}_{-10}(\text{Vf}, [s81], \text{K1}, \text{parameter}_{-1}, [s27], \text{K2}, \text{parameter}_{-1}, [s49], \text{K3},$$
parameter\_1, [s45], [s34], parameter\_1, K4, [s83], parameter\_1, K5, Vr, [s35]) (73)

 $function\_10 \left(Vf, r1, K1, n1, I1, K2, n2, I2, K3, n3, I3, A1, n4, K4, A2, n5, K5, Vr, p\right)$ 

$$= Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \frac{K3^{n3}}{I3^{n3} + K3^{n3}} \cdot \left(\frac{A1^{n4}}{A1^{n4} + K4^{n4}} + \frac{A2^{n5}}{A2^{n5} + K5^{n5}}\right) - Vr \cdot p$$
(74)

 $function\_10 \, (Vf, r1, K1, n1, I1, K2, n2, I2, K3, n3, I3, A1, n4, K4, A2, n5, K5, Vr, p)$ 

$$= Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \frac{K3^{n3}}{I3^{n3} + K3^{n3}} \cdot \left(\frac{A1^{n4}}{A1^{n4} + K4^{n4}} + \frac{A2^{n5}}{A2^{n5} + K5^{n5}}\right) - Vr \cdot p$$
(75)

Table 73: Properties of each parameter.

	14010 / 2	Troperties of each parameter.	
Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	lacksquare
K1	<b>K</b> 1	0.199	$\square$
K2	K2	9.615	$\square$
КЗ	K3	0.214	$\square$
K4	K4	0.321	
K5	K5	0.100	$\square$
Vr	Vr	0.100	

#### 9.18 Reaction re24

This is a reversible reaction of one reactant forming one product influenced by six modifiers.

Name re24

#### **Reaction equation**

$$s82 \stackrel{\text{s25, s33, s82, s25, s33, s34}}{\rightleftharpoons} s34$$
 (76)

#### Reactant

Table 74: Properties of each reactant.

Id	Name	SBO
s82	STAT6	

## **Modifiers**

Table 75: Properties of each modifier.

Id	Name	SBO
s25	IFNg-IFNgR	
s33	IL4-IL4R	
s82	STAT6	
s25	IFNg-IFNgR	
s33	IL4-IL4R	
s34	STAT6-P	

#### **Product**

Table 76: Properties of each product.

Id	Name	SBO
s34	STAT6-P	

## **Kinetic Law**

$$v_{18} = \text{vol}(c1) \cdot \text{function}_{15}(Vf, [s82], K, \text{parameter}_{1}, [s25], [s33], n1, K1, Vr, [s34])$$
 (77)

$$\text{function\_15} \left( Vf, r1, K, n, I, A1, n1, K1, Vr, p \right) = Vf \cdot r1 \cdot \frac{K^n}{I^n + K^n} \cdot \frac{A1^{n1}}{A1^{n1} + K1^{n1}} - Vr \cdot p \quad (78)$$

$$function\_15 \left(Vf, r1, K, n, I, A1, n1, K1, Vr, p\right) = Vf \cdot r1 \cdot \frac{K^n}{I^n + K^n} \cdot \frac{A1^{n1}}{A1^{n1} + K1^{n1}} - Vr \cdot p \quad (79)$$

Table 77: Properties of each parameter.

Tuble 17. I Toperties of each parameter.			
Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	
K	K	0.100	
n1	n1	0.004	
K1	<b>K</b> 1	0.100	
Vr	Vr	0.100	$\square$

# 9.19 Reaction re25

This is a reversible reaction of two reactants forming one product influenced by three modifiers.

Name re25

# **Reaction equation**

$$889 + 837 \xrightarrow{889, 837, 838} 838$$
 (80)

## **Reactants**

Table 78: Properties of each reactant.

Id	Name	SBO
s89	eIL2	
s37	IL2R	

## **Modifiers**

Table 79: Properties of each modifier.

		_	
	Id	Name	SBO
•	s37	eIL2 IL2R IL2-IL2R	

## **Product**

Table 80: Properties of each product.

Id	Name	SBO
s38	IL2-IL2R	

#### **Kinetic Law**

$$v_{19} = k1 \cdot [s89] \cdot [s37] - k2 \cdot [s38] \tag{81}$$

Table 81: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k1	k1	0.1	$\square$
k2	k2	0.1	$\square$

# 9.20 Reaction re26

This is a reversible reaction of one reactant forming one product influenced by four modifiers.

Name re26

# **Reaction equation**

$$s77 \stackrel{s38, s77, s38, s39}{\rightleftharpoons} s39$$
 (82)

#### Reactant

Table 82: Properties of each reactant.

Id	Name	SBO
s77	STAT5	

### **Modifiers**

Table 83: Properties of each modifier.

Id	Name	SBO
s38	IL2-IL2R	
s77	STAT5	
s38	IL2-IL2R	
s39	STAT5-P	

## **Product**

Table 84: Properties of each product.

Id	Name	SBO
s39	STAT5-P	

## **Kinetic Law**

#### **Derived unit** contains undeclared units

$$v_{20} = \text{vol}(c1) \cdot \text{function}_2(Vf, [s77], [s38], \text{parameter}_1, K, Vr, [s39])$$
 (83)

$$\text{function\_2}\left(Vf,r1,A,n,K,Vr,p\right) = Vf \cdot r1 \cdot \left(1 + \frac{A^n}{A^n + K^n}\right) - Vr \cdot p \tag{84}$$

$$function\_2\left(Vf,r1,A,n,K,Vr,p\right) = Vf \cdot r1 \cdot \left(1 + \frac{A^n}{A^n + K^n}\right) - Vr \cdot p \tag{85}$$

Table 85: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	$\checkmark$
K	K	0.375	
Vr	Vr	0.100	

#### 9.21 Reaction re27

This is a reversible reaction of one reactant forming one product influenced by 16 modifiers.

## Name re27

### **Reaction equation**

$$s78 = \frac{s34, s49, s48, s45, s26, s39, s83, s78, s34, s49, s48, s45, s26, s39, s83, s40}{s40}$$
 s40 (86)

#### Reactant

Table 86: Properties of each reactant.

Id	Name	SBO
s78	FOXP3	

#### **Modifiers**

Table 87: Properties of each modifier.

Id	Name	SBO
s34	STAT6-P	
s49	STAT3-P	
s48	IL6-IL6R	
s45	TGFb-TGFbR	
s26	STAT1-P	
s39	STAT5-P	
s83	L-PPARg	
s78	FOXP3	
s34	STAT6-P	
s49	STAT3-P	
s48	IL6-IL6R	
s45	TGFb-TGFbR	
s26	STAT1-P	
s39	STAT5-P	
s83	L-PPARg	
s40	acetylated FOXP3	

#### **Product**

Table 88: Properties of each product.

Id	Name	SBO
s40	acetylated FOXP3	

## **Kinetic Law**

$$v_{21} = vol\left(c1\right) \cdot function\_11\left(Vf, [s78], K1, parameter\_1, [s34], K2, parameter\_1, [s49], K3, \\ parameter\_1, [s48], [s45], parameter\_1, K4, [s26], parameter\_1, K5, [s39], parameter\_1, K6, \\ [s83], parameter\_1, K7, Vr, [s40]\right)$$
 (87)

$$\begin{split} &\text{function\_11}\left(Vf,r1,K1,n1,I1,K2,n2,I2,K3,n3,I3,A1,n4,K4,A2,n5,K5,A3,\\ &n6,K6,A4,n7,K7,Vr,p\right) = Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \frac{K3^{n3}}{I3^{n3} + K3^{n3}} \\ & \cdot \left(\frac{A1^{n4}}{A1^{n4} + K4^{n4}} + \frac{A2^{n5}}{A2^{n5} + K5^{n5}} + \frac{A3^{n6}}{A3^{n6} + K6^{n6}} + \frac{A4^{n7}}{A4^{n7} + K7^{n7}}\right) - Vr \cdot p \end{split} \tag{88}$$

$$n6, K6, A4, n7, K7, Vr, p) = Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \frac{K3^{n3}}{I3^{n3} + K3^{n3}}$$

$$\cdot \left( \frac{A1^{n4}}{A1^{n4} + K4^{n4}} + \frac{A2^{n5}}{A2^{n5} + K5^{n5}} + \frac{A3^{n6}}{A3^{n6} + K6^{n6}} + \frac{A4^{n7}}{A4^{n7} + K7^{n7}} \right) - Vr \cdot p$$
(89)

Table 89: Properties of each parameter.

Id	Name	SBO	Value	Unit	Constant
Vf	Vf		0.100		$ \overline{\checkmark} $
K1	K1		100.000		$\overline{Z}$
K2	K2		0.355		$\overline{\checkmark}$
КЗ	K3		1.313		$   \overline{\mathscr{L}} $
K4	K4	6	$5.79025 \cdot 10^{-4}$	ļ	$\mathbf{Z}$
K5	K5		2.079		$\mathbf{Z}$
K6	K6		100.000		
K7	K7	1	$.93254 \cdot 10^{-7}$	1	
Vr	Vr		0.100		

### 9.22 Reaction re28

This is a reversible reaction of two reactants forming one product influenced by three modifiers.

Name re28

## **Reaction equation**

$$887 + 844 \xrightarrow{887, 844, 845} 845$$
 (90)

#### **Reactants**

Table 90: Properties of each reactant.

Id	Name	SBO
s87 s44	eTGFb TGFbR	

# **Modifiers**

Table 91: Properties of each modifier.

Id	Name	SBO
s87	eTGFb	
s44	TGFbR	
s45	TGFb-TGFbR	

## **Product**

Table 92: Properties of each product.

Id	Name	SBO
s45	TGFb-TGFbR	

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{22} = k1 \cdot [s87] \cdot [s44] - k2 \cdot [s45]$$
 (91)

Table 93: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k1	k1	0.1	
k2	k2	0.1	$\checkmark$

# 9.23 Reaction re29

This is a reversible reaction of two reactants forming one product influenced by three modifiers.

#### Name re29

# **Reaction equation**

$$s90 + s47 \stackrel{\underline{s90, s47, s48}}{=} s48$$
 (92)

## **Reactants**

Table 94: Properties of each reactant.

Id	Name	SBO
200	eIL6 IL6R	

## **Modifiers**

Table 95: Properties of each modifier.

Id	Name	SBO
s90	eIL6	
s47	IL6R	
s48	IL6-IL6R	

### **Product**

Table 96: Properties of each product.

Id	Name	SBO
s48	IL6-IL6R	

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{23} = k1 \cdot [s90] \cdot [s47] - k2 \cdot [s48]$$
 (93)

Table 97: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k1	k1	0.1	$ \overline{\mathscr{A}} $
k2	k2	0.1	$\mathbf{Z}$

# 9.24 Reaction re30

This is a reversible reaction of one reactant forming one product influenced by twelve modifiers.

Name re30

# **Reaction equation**

$$s76 = \frac{s83, s54, s59, s63, s48, s76, s83, s54, s59, s63, s48, s49}{s49}$$
 (94)

## Reactant

Table 98: Properties of each reactant.

Id	Name	SBO
s76	STAT3	

# **Modifiers**

Table 99: Properties of each modifier.

Id	Name	SBO
s83	L-PPARg	
s54	IL21-IL21R	
s59	IL23-IL23R	
s63	IL10-IL10R	
s48	IL6-IL6R	
s76	STAT3	
s83	L-PPARg	
s54	IL21-IL21R	
s59	IL23-IL23R	
s63	IL10-IL10R	
s48	IL6-IL6R	
s49	STAT3-P	

# **Product**

Table 100: Properties of each product.

Id	Name	SBO
s49	STAT3-P	

#### **Kinetic Law**

$$\begin{array}{c} v_{24} = vol\left(c1\right) \cdot function\_12\left(Vf, [s76], K, parameter\_1, [s83], [s54], parameter\_1, K1, [s59], \\ parameter\_1, K2, [s63], parameter\_1, K3, [s48], parameter\_1, K4, Vr, [s49]) \end{array} \tag{95}$$

$$\begin{split} & \text{function\_12} \left( Vf, r1, K, n, I, A1, n1, K1, A2, n2, K2, A3, n3, K3, A4, n4, K4, Vr, p \right) \\ &= Vf \cdot r1 \cdot \frac{K^n}{I^n + K^n} \cdot \left( \frac{A1^{n1}}{A1^{n1} + K1^{n1}} + \frac{A2^{n2}}{A2^{n2} + K2^{n2}} + \frac{A3^{n3}}{A3^{n3} + K3^{n3}} + \frac{A4^{n4}}{A4^{n4} + K4^{n4}} \right) - Vr \cdot p \end{split} \tag{96}$$

$$function\_12 \left(Vf, r1, K, n, I, A1, n1, K1, A2, n2, K2, A3, n3, K3, A4, n4, K4, Vr, p\right)$$

$$= Vf \cdot r1 \cdot \frac{K^{n}}{I^{n} + K^{n}} \cdot \left(\frac{A1^{n1}}{A1^{n1} + K1^{n1}} + \frac{A2^{n2}}{A2^{n2} + K2^{n2}} + \frac{A3^{n3}}{A3^{n3} + K3^{n3}} + \frac{A4^{n4}}{A4^{n4} + K4^{n4}}\right) - Vr \cdot p$$
(97)

Table 101: Properties of each parameter.

Id         Name         SBO         Value         Unit         Constant           Vf         Vf         0.100         ✓           K         K         0.100         ✓           K1         K1         0.637         ✓           K2         K2         39.018         ✓           K3         K3         2.270         ✓           K4         K4         0.138         ✓           Vr         Vr         0.100         ✓			1	
K       K       0.100       ✓         K1       K1       0.637       ✓         K2       K2       39.018       ✓         K3       K3       2.270       ✓         K4       K4       0.138       ✓	Id	Name	SBO Value Unit	Constant
K1       K1       0.637         K2       K2       39.018         K3       K3       2.270         K4       K4       0.138	Vf	Vf	0.100	$   \sqrt{} $
K2       K2       39.018         K3       K3       2.270         K4       K4       0.138	K	K	0.100	
K3 K3 2.270   K4 K4 0.138   ✓	K1	K1	0.637	
K4 K4 0.138	K2	K2	39.018	
	КЗ	K3	2.270	$\overline{\mathbf{Z}}$
Vr Vr 0.100 ✓	K4	<b>K</b> 4	0.138	$\overline{\mathbf{Z}}$
<del>_</del>	Vr	Vr	0.100	

### 9.25 Reaction re31

This is a reversible reaction of one reactant forming one product influenced by ten modifiers.

Name re31

#### **Reaction equation**

$$s75 = \frac{s40, s83, s49, s45, s75, s40, s83, s49, s45, s50}{s50}$$
 s50 (98)

#### Reactant

Table 102: Properties of each reactant.

Id	Name	SBO
s75	RORgt	

#### **Modifiers**

Table 103: Properties of each modifier.

Id	Name	SBO
s40	acetylated FOXP3	
s83	L-PPARg	
s49	STAT3-P	
s45	TGFb-TGFbR	
s75	RORgt	
s40	acetylated FOXP3	
s83	L-PPARg	
s49	STAT3-P	
s45	TGFb-TGFbR	
<b>s</b> 50	RORgt-ligand	

#### **Product**

Table 104: Properties of each product.

Id	Name	SBO
s50	RORgt-ligand	

#### **Kinetic Law**

$$v_{25} = vol(c1) \cdot function\_5(Vf, [s75], K1, parameter\_1, [s40], K2, parameter\_1, [s83], [s49], parameter\_1, K3, [s45], parameter\_1, K4, Vr, [s50])$$

$$(99)$$

$$\begin{split} &\text{function\_5} \left( Vf, r1, K1, n1, I1, K2, n2, I2, A1, n3, K3, A2, n4, K4, Vr, p \right) \\ &= Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \left( \frac{A1^{n3}}{A1^{n3} + K3^{n3}} + \frac{A2^{n4}}{A2^{n4} + K4^{n4}} \right) - Vr \cdot p \end{split} \tag{100}$$

$$\begin{split} &\text{function\_5} \left( Vf, r1, K1, n1, I1, K2, n2, I2, A1, n3, K3, A2, n4, K4, Vr, p \right) \\ &= Vf \cdot r1 \cdot \frac{K1^{n1}}{I1^{n1} + K1^{n1}} \cdot \frac{K2^{n2}}{I2^{n2} + K2^{n2}} \cdot \left( \frac{A1^{n3}}{A1^{n3} + K3^{n3}} + \frac{A2^{n4}}{A2^{n4} + K4^{n4}} \right) - Vr \cdot p \end{split} \tag{101}$$

Table 105: Properties of each parameter.

Id	Name	SBO Value	Unit	Constant
Vf	Vf	0.225		
K1	K1	9722.090		
K2	K2	0.704		
КЗ	K3	1.241		$ \mathbf{Z} $
K4	K4	997.263		
Vr	Vr	0.100		

# 9.26 Reaction re32

This is a reversible reaction of two reactants forming one product influenced by seven modifiers.

#### Name re32

# **Reaction equation**

$$s51 + s53 \xrightarrow{s39, \text{ species\_3}, s51, s53, s39, \text{ species\_3}, s54} s54$$
 (102)

#### **Reactants**

Table 106: Properties of each reactant.

Id	Name	SBO
	eIL21 IL21R	

# **Modifiers**

Table 107: Properties of each modifier.

Id	Name	SBO
s39	STAT5-P	
species_3	IL17-IL17R	
s51	eIL21	
s53	IL21R	
s39	STAT5-P	
species_3	IL17-IL17R	
s54	IL21-IL21R	

## **Product**

Table 108: Properties of each product.

Id	Name	SBO
s54	IL21-IL21R	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$\nu_{26} = function\_14 \, (Vf, [s51], [s53], K, parameter\_1, [s39], [species\_3], parameter\_1, K1, Vr, [s54]) \eqno(103)$$

$$function\_14\left(Vf,r1,r2,K,n,I,A,n1,K1,Vr,p\right) = Vf \cdot r1 \cdot r2 \cdot \frac{K^{n}}{I^{n} + K^{n}} \cdot \left(1 + \frac{A^{n1}}{A^{n1} + K1^{n1}}\right) - Vr \cdot p \tag{104}$$

Table 109: Properties of each parameter.

		* *	
Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	
K	K	0.241	
K1	<b>K</b> 1	8.142	$\square$
Vr	Vr	0.100	

## 9.27 Reaction re33

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

Name re33

# **Reaction equation**

$$s52 \xrightarrow{s52} s51 \tag{105}$$

## Reactant

Table 110: Properties of each reactant.

Id	Name	SBO
s52	IL21	

# **Modifier**

Table 111: Properties of each modifier.

Id	Name	SBO
s52	IL21	

# **Product**

Table 112: Properties of each product.

Id	Name	SBO
s51	eIL21	

## **Kinetic Law**

Derived unit contains undeclared units

$$v_{27} = k1 \cdot [s52] \tag{106}$$

Table 113: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k1	k1	0.1	$\overline{\hspace{1cm}}$

# 9.28 Reaction re34

This is a reversible reaction of one reactant forming one product influenced by two modifiers.

Name re34

# **Reaction equation**

$$s55 \xrightarrow{s55, s57} s57 \tag{107}$$

## Reactant

Table 114: Properties of each reactant.

Id	Name	SBO
s55	eIL23	

## **Modifiers**

Table 115: Properties of each modifier.

Id	Name	SBO
s55	eIL23	
s57	p40/p19 dimer	

## **Product**

Table 116: Properties of each product.

Id	Name	SBO
s57	p40/p19 dimer	

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{28} = k1 \cdot [s55] - k2 \cdot [s57] \tag{108}$$

Table 117: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k1	k1	0.1	
k2	k2	0.1	$\square$

## 9.29 Reaction re35

This is a reversible reaction of two reactants forming one product influenced by five modifiers.

Name re35

### **Reaction equation**

$$s57 + s58 = \frac{s27, s58, s27, s59}{500} s59$$
 (109)

### **Reactants**

Table 118: Properties of each reactant.

Id	Name	SBO
s57	p40/p19 dimer	
s58	IL23R	

## **Modifiers**

Table 119: Properties of each modifier.

Id	Name	SBO
s27	Tbet-P	
s57	p40/p19 dimer	
s58	IL23R	
s27	Tbet-P	
s59	IL23-IL23R	

## **Product**

Table 120: Properties of each product.

Id	Name	SBO
s59	IL23-IL23R	

### **Kinetic Law**

$$v_{29} = vol(c1) \cdot function_1(Vf, [s57], [s58], K, parameter_1, [s27], Vr, [s59])$$
 (110)

$$function\_1\left(Vf,r1,r2,K,n,I,Vr,p\right) = Vf\cdot r1\cdot r2\cdot \frac{K^n}{I^n+K^n} - Vr\cdot p \tag{111}$$

$$\text{function\_1}\left(Vf,r1,r2,K,n,I,Vr,p\right) = Vf \cdot r1 \cdot r2 \cdot \frac{K^n}{I^n + K^n} - Vr \cdot p \tag{112}$$

Table 121: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	
K	K	4.661	
Vr	Vr	0.100	

# 9.30 Reaction re36

This is a reversible reaction of two reactants forming one product influenced by seven modifiers.

Name re36

# **Reaction equation**

$$species\_9 + species\_8 \xrightarrow{s83, s48, species\_9, species\_8, s83, s48, s57} s57$$
 (113)

#### **Reactants**

Table 122: Properties of each reactant.

Id	Name	SBO
species_9 species_8	p40 p19	

#### **Modifiers**

Table 123: Properties of each modifier.

Id	Name	SBO
s83	L-PPARg	
s48	IL6-IL6R	
species_9	p40	
species_8	p19	
s83	L-PPARg	
s48	IL6-IL6R	
s57	p40/p19 dimer	

### **Product**

Table 124: Properties of each product.

Id	Name	SBO
s57	p40/p19 dimer	

#### **Kinetic Law**

#### Derived unit contains undeclared units

$$v_{30} = \text{vol}(c1) \cdot \text{function\_14}(\text{Vf}, [\text{species\_9}], [\text{species\_8}], \text{K}, \text{parameter\_1}, [\text{s83}], [\text{s48}],$$

$$parameter\_1, \text{K1}, \text{Vr}, [\text{s57}])$$
(114)

$$function\_14\left(Vf,r1,r2,K,n,I,A,n1,K1,Vr,p\right) = Vf \cdot r1 \cdot r2 \cdot \frac{K^n}{I^n + K^n} \cdot \left(1 + \frac{A^{n1}}{A^{n1} + K1^{n1}}\right) - Vr \cdot p \tag{115}$$

$$function\_14 \left(Vf, r1, r2, K, n, I, A, n1, K1, Vr, p\right) = Vf \cdot r1 \cdot r2 \cdot \frac{K^n}{I^n + K^n} \cdot \left(1 + \frac{A^{n1}}{A^{n1} + K1^{n1}}\right) - Vr \cdot p \tag{116}$$

Table 125: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	$\overline{Z}$
K	K	0.100	
K1	K1	25.535	
Vr	Vr	0.100	$\checkmark$

# 9.31 Reaction re37

This is a reversible reaction of one reactant forming one product influenced by four modifiers.

Name re37

### **Reaction equation**

species\_6 
$$\stackrel{\text{s49, species_6, s49, s52}}{=\!=\!=\!=\!=}$$
 s52 (117)

## Reactant

Table 126: Properties of each reactant.

Id	Name	SBO
species_6	pIL21	

## **Modifiers**

Table 127: Properties of each modifier.

Id	Name	SBO
s49	STAT3-P	
species_6	pIL21	
s49	STAT3-P	
s52	IL21	

#### **Product**

Table 128: Properties of each product.

Id	Name	SBO
s52	IL21	

#### **Kinetic Law**

$$v_{31} = \text{vol}(c1) \cdot \text{function} 2 (Vf, [\text{species\_6}], [\text{s49}], \text{parameter\_1}, K, Vr, [\text{s52}])$$
 (118)

$$\text{function\_2}\left(Vf,r1,A,n,K,Vr,p\right) = Vf \cdot r1 \cdot \left(1 + \frac{A^n}{A^n + K^n}\right) - Vr \cdot p \tag{119}$$

$$function\_2\left(Vf,r1,A,n,K,Vr,p\right) = Vf \cdot r1 \cdot \left(1 + \frac{A^n}{A^n + K^n}\right) - Vr \cdot p \tag{120}$$

Table 129: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	Ø
K	K	0.119	
Vr	Vr	0.100	$\checkmark$

# 9.32 Reaction re39

This is a reversible reaction of two reactants forming one product influenced by three modifiers.

Name re38

# **Reaction equation**

species\_2 + s62 
$$\stackrel{\text{species}\_2, s62, s63}{\rightleftharpoons}$$
 s63 (121)

#### **Reactants**

Table 130: Properties of each reactant.

Id	Name	SBO
species_2 s62	eIL10 IL10R	

#### **Modifiers**

Table 131: Properties of each modifier.

Id	Name	SBO
species_2 s62 s63	eIL10 IL10R IL10-IL10R	

## **Product**

Table 132: Properties of each product.

Id	Name	SBO
s63	IL10-IL10R	

# **Kinetic Law**

$$v_{32} = k1 \cdot [\text{species}_2] \cdot [\text{s}62] - k2 \cdot [\text{s}63]$$
 (122)

Table 133: Properties of each parameter.

		1 1	
Id	Name	SBO Value Unit	Constant
k1	k1	0.1	
k2	k2	0.1	

## 9.33 Reaction re42

This is a reversible reaction of two reactants forming one product influenced by seven modifiers.

Name re42

# **Reaction equation**

$$s85 + s86 \xrightarrow{s25, s33, s85, s86, s25, s33, s83} s83$$
 (123)

#### **Reactants**

Table 134: Properties of each reactant.

Id	Name	SBO
s85	PPARg	
s86	Ligand	

## **Modifiers**

Table 135: Properties of each modifier.

Id	Name	SBO
s25	IFNg-IFNgR	
s33	IL4-IL4R	
s85	PPARg	
s86	Ligand	
s25	IFNg-IFNgR	
s33	IL4-IL4R	
s83	L-PPARg	

#### **Product**

Table 136: Properties of each product.

Id	Name	SBO
s83	L-PPARg	

#### **Kinetic Law**

#### **Derived unit** contains undeclared units

$$v_{33} = \text{vol}(c1) \cdot \text{function}_14(\text{Vf}, [s85], [s86], \text{K}, \text{parameter}_1, [s25], [s33], \text{parameter}_1, \\ \text{K1}, \text{Vr}, [s83])$$
 (124)

$$function_{-}14\left(Vf,r1,r2,K,n,I,A,n1,K1,Vr,p\right) = Vf \cdot r1 \cdot r2 \cdot \frac{K^{n}}{I^{n}+K^{n}} \cdot \left(1 + \frac{A^{n1}}{A^{n1}+K1^{n1}}\right) - Vr \cdot p \tag{125}$$

$$function_{-}14\left(Vf,r1,r2,K,n,I,A,n1,K1,Vr,p\right) = Vf \cdot r1 \cdot r2 \cdot \frac{K^{n}}{I^{n} + K^{n}} \cdot \left(1 + \frac{A^{n1}}{A^{n1} + K1^{n1}}\right) - Vr \cdot p \tag{126}$$

Table 137: Properties of each parameter.

		1	
Id	Name	SBO Value Unit	Constant
Vf	Vf	0.1	$\overline{Z}$
K	K	0.1	$\square$
K1	K1	0.1	
Vr	Vr	0.1	$\square$

#### 9.34 Reaction re44

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

Name re44

## **Reaction equation**

$$s43 \xrightarrow{s43} s87 \tag{127}$$

#### Reactant

Table 138: Properties of each reactant.

Id	Name	SBO
s43	TGFb	

## **Modifier**

Table 139: Properties of each modifier.

Id	Name	SBO
s43	TGFb	

# **Product**

Table 140: Properties of each product.

Id	Name	SBO
s87	eTGFb	

## **Kinetic Law**

Derived unit contains undeclared units

$$v_{34} = k1 \cdot [s43] \tag{128}$$

Table 141: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k1	k1	0.1	$\overline{\hspace{1cm}}$

## 9.35 Reaction re45

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

# Name re45

# **Reaction equation**

$$s36 \xrightarrow{s36} s89 \tag{129}$$

## Reactant

Table 142: Properties of each reactant.

Id	Name	SBO
s36	IL2	

## Modifier

Table 143: Properties of each modifier.

Id	Name	SBO
s36	IL2	

## **Product**

Table 144: Properties of each product.

Id	Name	SBO
s89	eIL2	

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{35} = k1 \cdot [s36] \tag{130}$$

Table 145: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k1	k1	0.1	

## 9.36 Reaction re46

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

# Name re46

# **Reaction equation**

$$s46 \xrightarrow{s46} s90 \tag{131}$$

#### Reactant

Table 146: Properties of each reactant.

Id	Name	SBO
s46	IL6	

#### **Modifier**

Table 147: Properties of each modifier.

Id	Name	SBO
s46	IL6	

#### **Product**

Table 148: Properties of each product.

Id	Name	SBO
s90	eIL6	

## **Kinetic Law**

Derived unit contains undeclared units

$$v_{36} = k1 \cdot [s46] \tag{132}$$

Table 149: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k1	k1	0.1	

# 9.37 Reaction reaction\_1

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

# Name re48

# **Reaction equation**

$$s74 \xrightarrow{s74} species_2$$
 (133)

## Reactant

Table 150: Properties of each reactant.

Id	Name	SBO
s74	IL10	

# **Modifier**

Table 151: Properties of each modifier.

Id	Name	SBO
s74	IL10	

# **Product**

Table 152: Properties of each product.

Id	Name	SBO
species_2	eIL10	

# **Kinetic Law**

$$v_{37} = k1 \cdot [s74] \tag{134}$$

Table 153: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k1	k1	0.1	

# 9.38 Reaction reaction\_2

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

Name re49

# **Reaction equation**

$$s73 \xrightarrow{s73} species_1$$
 (135)

#### Reactant

Table 154: Properties of each reactant.

Id	Name	SBO
s73	IL17	

#### **Modifier**

Table 155: Properties of each modifier.

Id	Name	SBO
s73	IL17	

#### **Product**

Table 156: Properties of each product.

Id	Name	SBO
species_1	eIL17	

## **Kinetic Law**

$$v_{38} = k1 \cdot [s73] \tag{136}$$

Table 157: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k1	k1	0.1	

# 9.39 Reaction reaction\_3

This is a reversible reaction of two reactants forming one product influenced by three modifiers.

Name re50

# **Reaction equation**

$$species_1 + species_4 \xrightarrow{species_1, species_4, species_3} species_3$$
 (137)

#### **Reactants**

Table 158: Properties of each reactant.

Id	Name	SBO
species_1 species_4		

#### **Modifiers**

Table 159: Properties of each modifier.

Id	Name	SBO
species_1	eIL17	
${ t species\_4}$	IL17R	
species_3	IL17-IL17R	

#### **Product**

Table 160: Properties of each product.

Id	Name	SBO
species_3	IL17-IL17R	

## **Kinetic Law**

$$v_{39} = k1 \cdot [\text{species\_1}] \cdot [\text{species\_4}] - k2 \cdot [\text{species\_3}]$$
 (138)

Table 161: Properties of each parameter.

т.і	Nama	CDO Volum IImia	Canatant
10	Name	SBO Value Unit	Constant
k1	k1	0.185	$\square$
k2	k2	0.190	

## 9.40 Reaction reaction\_4

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

Name re47

# **Reaction equation**

$$s1 \xrightarrow{s1} species_12$$
 (139)

## Reactant

Table 162: Properties of each reactant.

Id	Name	SBO
s1	IL18	

#### **Modifier**

Table 163: Properties of each modifier.

Id	Name	SBO
s1	IL18	

## **Product**

Table 164: Properties of each product.

Id	Name	SBO
species_12	eIL18	

#### **Kinetic Law**

$$v_{40} = \mathbf{k}1 \cdot [\mathbf{s}1] \tag{140}$$

Table 165: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
k1	k1	0.1	

## 9.41 Reaction reaction\_5

This is a reversible reaction of one reactant forming one product influenced by ten modifiers.

### Name re51

## **Reaction equation**

species\_7 
$$\stackrel{$83, $59, $50, $21, $pecies_7, $83, $59, $50, $21, $73}{=} $73$$
 (141)

### Reactant

Table 166: Properties of each reactant.

Id	Name	SBO
species_7	pIL17	

#### **Modifiers**

Table 167: Properties of each modifier.

Id	Name	SBO
s83	L-PPARg	
s59	IL23-IL23R	
<b>s</b> 50	RORgt-ligand	
s21	STAT4-P	
species_7	pIL17	
s83	L-PPARg	
s59	IL23-IL23R	
s50	RORgt-ligand	
s21	STAT4-P	
s73	IL17	

#### **Product**

Table 168: Properties of each product.

Id	Name	SBO
s73	IL17	

#### **Kinetic Law**

#### Derived unit contains undeclared units

$$v_{41} = vol(c1) \cdot function\_6(Vf, [species\_7], K, parameter\_1, [s83], [s59], parameter\_1, K1, [s50], parameter\_1, K2, [s21], parameter\_1, K3, Vr, [s73])$$
 (142)

$$\begin{split} &\text{function\_6}\left(Vf, r1, K, n, I, A1, n1, K1, A2, n2, K2, A3, n3, K3, Vr, p\right) \\ &= Vf \cdot r1 \cdot \frac{K^n}{I^n + K^n} \cdot \left(\frac{A1^{n1}}{A1^{n1} + K1^{n1}} + \frac{A2^{n2}}{A2^{n2} + K2^{n2}} + \frac{A3^{n3}}{A3^{n3} + K3^{n3}}\right) - Vr \cdot p \end{split} \tag{143}$$

$$\begin{split} &\text{function\_6}\left(Vf,r1,K,n,I,A1,n1,K1,A2,n2,K2,A3,n3,K3,Vr,p\right) \\ &= Vf \cdot r1 \cdot \frac{K^n}{I^n + K^n} \cdot \left(\frac{A1^{n1}}{A1^{n1} + K1^{n1}} + \frac{A2^{n2}}{A2^{n2} + K2^{n2}} + \frac{A3^{n3}}{A3^{n3} + K3^{n3}}\right) - Vr \cdot p \end{split} \tag{144}$$

Table 169: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
Vf	Vf	0.225	$\checkmark$
K	K	0.100	
K1	<b>K</b> 1	1.629	
K2	K2	0.527	
КЗ	K3	5.479	
Vr	Vr	0.100	$\overline{\mathbf{Z}}$

## 9.42 Reaction reaction\_6

This is a reversible reaction of one reactant forming one product influenced by two modifiers.

Name IL18 pool

#### **Reaction equation**

$$species_{16} = \frac{species_{16}, species_{12}}{species_{12}}$$
 species\_12 (145)

### Reactant

Table 170: Properties of each reactant.

Id	Name	SBO
species_16	IL18_pool	

#### **Modifiers**

Table 171: Properties of each modifier.

Id	Name	SBO
species_16 species_12	-	

### **Product**

Table 172: Properties of each product.

Id	Name	SBO
species_12	eIL18	

### **Kinetic Law**

$$v_{42} = \text{vol}(\text{default}) \cdot \text{function\_16}(V, [\text{species\_16}], \text{parameter\_1}, [\text{species\_12}], k)$$
 (146)

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{147}$$

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{148}$$

Table 173: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
V	V	0.1	$ \overline{\checkmark} $
k	k	0.5	$\overline{\checkmark}$

### 9.43 Reaction reaction\_7

This is a reversible reaction of one reactant forming one product influenced by two modifiers.

Name IL12 pool

## **Reaction equation**

species\_17 
$$\rightleftharpoons$$
 s11 (149)

#### Reactant

Table 174: Properties of each reactant.

Id	Name	SBO
species_17	IL12_pool	

#### **Modifiers**

Table 175: Properties of each modifier.

Id	Name	SBO
species_17 s11	IL12_pool eIL12	

#### **Product**

Table 176: Properties of each product.

Id	Name	SBO
s11	eIL12	

#### **Kinetic Law**

$$v_{43} = \text{vol}(\text{default}) \cdot \text{function\_16}(V, [\text{species\_17}], \text{parameter\_1}, [\text{s11}], k)$$
 (150)

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{151}$$

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{152}$$

Table 177: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
V	V	0.1	
k	k	0.5	$\checkmark$

### 9.44 Reaction reaction\_8

This is a reversible reaction of one reactant forming one product influenced by two modifiers.

Name IFNg pool

## **Reaction equation**

species\_18 
$$\stackrel{\text{species}\_18, s22}{\longleftarrow}$$
 s22 (153)

#### Reactant

Table 178: Properties of each reactant.

Id	Name	SBO
species_18	IFNg_pool	

#### **Modifiers**

Table 179: Properties of each modifier.

Id	Name	SBO
species_18 s22	IFNg_pool eIFNg	

### **Product**

Table 180: Properties of each product.

Id	Name	SBO
s22	eIFNg	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{44} = \text{vol}(\text{default}) \cdot \text{function\_16}(V, [\text{species\_18}], \text{parameter\_1}, [\text{s22}], k)$$
 (154)

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{155}$$

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{156}$$

Table 181: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
V	V	0.1	
k	k	0.1	

### 9.45 Reaction reaction\_9

This is a reversible reaction of one reactant forming one product influenced by two modifiers.

Name IL21 pool

## **Reaction equation**

species\_19 
$$\stackrel{\text{species}\_19, s51}{\longleftarrow}$$
 s51 (157)

#### Reactant

Table 182: Properties of each reactant.

Id	Name	SBO
species_19	IL21_pool	

#### **Modifiers**

Table 183: Properties of each modifier.

Id	Name	SBO
species_19 s51	IL21_pool eIL21	

#### **Product**

Table 184: Properties of each product.

Id	Name	SBO
s51	eIL21	

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{45} = \text{vol}(\text{default}) \cdot \text{function}_{16}(V, [\text{species}_{19}], \text{parameter}_{1}, [\text{s51}], k)$$
 (158)

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{159}$$

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{160}$$

Table 185: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
V	V	0.1	
k	k	0.1	$\checkmark$

#### 9.46 Reaction reaction\_10

This is a reversible reaction of one reactant forming one product influenced by two modifiers.

Name IL23 pool

### **Reaction equation**

species\_20 
$$\stackrel{\text{species}\_20, s55}{=\!=\!=\!=\!=} s55$$
 (161)

## Reactant

Table 186: Properties of each reactant.

Id	Name	SBO
species_20	IL23_pool	

#### **Modifiers**

Table 187: Properties of each modifier.

Id	Name	SBO
species_20 s55	IL23_pool eIL23	

### **Product**

Table 188: Properties of each product.

Id	Name	SBO
s55	eIL23	

### **Kinetic Law**

$$v_{46} = \text{vol}(\text{default}) \cdot \text{function\_16}(V, [\text{species\_20}], \text{parameter\_1}, [\text{s55}], k)$$
 (162)

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{163}$$

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{164}$$

Table 189: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
V	V	0.1	
k	k	0.1	

### 9.47 Reaction reaction\_11

This is a reversible reaction of one reactant forming one product influenced by two modifiers.

Name IL17 pool

## **Reaction equation**

species\_21 
$$\rightleftharpoons$$
 s73 s73 (165)

#### Reactant

Table 190: Properties of each reactant.

Id	Name	SBO
species_21	IL17_pool	

#### **Modifiers**

Table 191: Properties of each modifier.

Id	Name	SBO
species_21	IL17_pool	
s73	IL17	

#### **Product**

Table 192: Properties of each product.

Id	Name	SBO
s73	IL17	

### **Kinetic Law**

$$v_{47} = \text{function}_{16}(V, [\text{species}_{21}], \text{parameter}_{1}, [\text{s}73], k)$$
 (166)

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{167}$$

Table 193: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
٧	V	0.1	$\square$
k	k	0.1	

### 9.48 Reaction reaction\_12

This is a reversible reaction of one reactant forming one product influenced by two modifiers.

Name IL10 pool

## **Reaction equation**

### Reactant

Table 194: Properties of each reactant.

Id	Name	SBO
species_22	IL10_pool	

#### **Modifiers**

Table 195: Properties of each modifier.

Id	Name	SBO
species_22	IL10_pool	
${\tt species\_2}$	eIL10	

## **Product**

Table 196: Properties of each product.

Id	Name	SBO
species_2	eIL10	

#### **Kinetic Law**

$$v_{48} = \text{vol}(\text{default}) \cdot \text{function\_16}(V, [\text{species\_22}], \text{parameter\_1}, [\text{species\_2}], k)$$
 (169)

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{170}$$

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{171}$$

Table 197: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
V	V	0.1	$\overline{Z}$
k	k	0.1	

### 9.49 Reaction reaction\_13

This is a reversible reaction of one reactant forming one product influenced by two modifiers.

Name IL6 pool

### **Reaction equation**

species\_23 
$$\stackrel{\text{species}\_23, s90}{\longleftarrow}$$
 s90 (172)

#### Reactant

Table 198: Properties of each reactant.

Id	Name	SBO
species_23	IL6_pool	

### **Modifiers**

Table 199: Properties of each modifier.

Id	Name	SBO
species_23	-	
s90	eIL6	

## **Product**

Table 200: Properties of each product.

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{49} = \text{vol}(\text{default}) \cdot \text{function\_16}(V, [\text{species\_23}], \text{parameter\_1}, [\text{s90}], k)$$
 (173)

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{174}$$

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{175}$$

Table 201: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
V	V	0.1	$\overline{\checkmark}$
k	k	0.1	$\checkmark$

### 9.50 Reaction reaction\_14

This is a reversible reaction of one reactant forming one product influenced by two modifiers.

Name IL2 pool

### **Reaction equation**

$$species_24 \xrightarrow{species_24, s89} s89 \tag{176}$$

#### Reactant

Table 202: Properties of each reactant.

Id	Name	SBO
species_24	IL2_pool	

### **Modifiers**

Table 203: Properties of each modifier.

Id	Name	SBO
species_24 s89	IL2_pool eIL2	

### **Product**

Table 204: Properties of each product.

Id	Name	SBO
s89	eIL2	

### **Kinetic Law**

$$v_{50} = \text{vol}(\text{default}) \cdot \text{function\_16}(V, [\text{species\_24}], \text{parameter\_1}, [\text{s89}], k)$$
 (177)

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{178}$$

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{179}$$

Table 205: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
V	V	0.1	
k	k	0.1	

### 9.51 Reaction reaction\_15

This is a reversible reaction of one reactant forming one product influenced by two modifiers.

Name TGFb pool

## **Reaction equation**

species\_25 
$$\rightleftharpoons$$
 second second second species\_25, s87 (180)

#### Reactant

Table 206: Properties of each reactant.

Id	Name	SBO
species_25	TGFb_pool	

#### **Modifiers**

Table 207: Properties of each modifier.

Id	Name	SBO
species_25 s87	TGFb_pool eTGFb	

#### **Product**

Table 208: Properties of each product.

Id	Name	SBO
s87	eTGFb	

#### **Kinetic Law**

$$v_{51} = \text{vol}(\text{default}) \cdot \text{function\_16}(V, [\text{species\_25}], \text{parameter\_1}, [\text{s87}], k)$$
 (181)

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{182}$$

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{183}$$

Table 209: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
V	V	0.1	
k	k	0.1	

### 9.52 Reaction reaction\_16

This is a reversible reaction of one reactant forming one product influenced by two modifiers.

Name IL4 pool

## **Reaction equation**

species 
$$26 \rightleftharpoons 30$$
 species  $30 \rightleftharpoons 30$  (184)

#### Reactant

Table 210: Properties of each reactant.

Id	Name	SBO
species_26	IL4_pool	

#### **Modifiers**

Table 211: Properties of each modifier.

Id	Name	SBO
species_26 s30	IL4_pool eIL4	

## **Product**

Table 212: Properties of each product.

Id	Name	SBO
s30	eIL4	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{52} = \text{vol}(\text{default}) \cdot \text{function\_16}(V, [\text{species\_26}], \text{parameter\_1}, [\text{s30}], k)$$
 (185)

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{186}$$

$$function\_16\left(V,pool,n,ext,k\right) = V \cdot \left(\frac{pool^n}{pool^n + ext^n + 0.0010} - k \cdot ext\right) \tag{187}$$

Table 213: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
V	V	0.1	$\overline{Z}$
k	k	0.1	$\checkmark$

#### 9.53 Reaction reaction\_17

This is a reversible reaction of one reactant forming one product influenced by twelve modifiers.

#### Name re52

## **Reaction equation**

$$species_27 \xrightarrow{s59, s83, s54, s35, s39, species_27, s59, s83, s54, s35, s39, s74} s74$$
 (188)

#### Reactant

Table 214: Properties of each reactant.

Id	Name	SBO
species_27	pIL10	

#### **Modifiers**

Table 215: Properties of each modifier.

Id	Name	SBO
s59	IL23-IL23R	
s83	L-PPARg	
s54	IL21-IL21R	
s35	GATA3-P	
s39	STAT5-P	
species_27	pIL10	
s59	IL23-IL23R	
s83	L-PPARg	
s54	IL21-IL21R	
s35	GATA3-P	
s39	STAT5-P	
s74	IL10	

#### **Product**

Table 216: Properties of each product.

Id	Name	SBO
s74	IL10	

#### **Kinetic Law**

$$\begin{split} & \text{function\_12} \left( Vf, r1, K, n, I, A1, n1, K1, A2, n2, K2, A3, n3, K3, A4, n4, K4, Vr, p \right) \\ & = Vf \cdot r1 \cdot \frac{K^n}{I^n + K^n} \cdot \left( \frac{A1^{n1}}{A1^{n1} + K1^{n1}} + \frac{A2^{n2}}{A2^{n2} + K2^{n2}} + \frac{A3^{n3}}{A3^{n3} + K3^{n3}} + \frac{A4^{n4}}{A4^{n4} + K4^{n4}} \right) - Vr \cdot p \end{split} \tag{190}$$

Table 217: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
Vf	Vf	0.100	$lue{oldsymbol{arphi}}$
K	K	0.508	<b>Z</b>

Id	Name	SBO Value Unit	Constant
K1	K1	0.100	$\overline{Z}$
K2	K2	0.001	$   \overline{\checkmark} $
КЗ	K3	0.645	
K4	K4	100.000	$   \overline{\checkmark} $
Vr	Vr	0.100	$   \overline{\checkmark} $

# 10 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.

### **10.1 Species** s22

Name eIFNg

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

Initial assignment s22

This species takes part in five reactions (as a reactant in re11 and as a product in re12, reaction—8 and as a modifier in re11, reaction—8).

$$\frac{\mathrm{d}}{\mathrm{d}t}s22 = |v_8| + |v_{44}| - |v_7| \tag{191}$$

### **10.2 Species** s11

Name eIL12

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

Initial assignment s11

This species takes part in five reactions (as a reactant in re8 and as a product in re6, reaction—7 and as a modifier in re8, reaction—7).

$$\frac{\mathrm{d}}{\mathrm{d}t}s11 = |v_3| + |v_{43}| - |v_4| \tag{192}$$

## **10.3 Species** s51

Name eIL21

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

#### Initial assignment s51

This species takes part in five reactions (as a reactant in re32 and as a product in re33, reaction—9 and as a modifier in re32, reaction—9).

$$\frac{\mathrm{d}}{\mathrm{d}t}s51 = |v_{27}| + |v_{45}| - |v_{26}| \tag{193}$$

## **10.4 Species** s55

Name eIL23

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

#### Initial assignment s55

This species takes part in four reactions (as a reactant in re34 and as a product in reaction\_10 and as a modifier in re34, reaction\_10).

$$\frac{d}{dt}s55 = |v_{46}| - |v_{28}| \tag{194}$$

#### **10.5 Species** s30

Name eIL4

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

### Initial assignment s30

This species takes part in five reactions (as a reactant in re19 and as a product in re18, reaction—16 and as a modifier in re19, reaction—16).

$$\frac{\mathrm{d}}{\mathrm{d}t}s30 = |v_{14}| + |v_{52}| - |v_{15}| \tag{195}$$

### **10.6 Species** s87

Name eTGFb

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

### Initial assignment s87

This species takes part in five reactions (as a reactant in re28 and as a product in re44, reaction\_15 and as a modifier in re28, reaction\_15).

$$\frac{\mathrm{d}}{\mathrm{d}t} s87 = |v_{34}| + |v_{51}| - |v_{22}| \tag{196}$$

## **10.7 Species** s89

Name eIL2

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

#### Initial assignment s89

This species takes part in five reactions (as a reactant in re25 and as a product in re45, reaction—14 and as a modifier in re25, reaction—14).

$$\frac{\mathrm{d}}{\mathrm{d}t}s89 = |v_{35}| + |v_{50}| - |v_{19}| \tag{197}$$

## **10.8 Species** s90

Name eIL6

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

#### Initial assignment s90

This species takes part in five reactions (as a reactant in re29 and as a product in re46, reaction—13 and as a modifier in re29, reaction—13).

$$\frac{\mathrm{d}}{\mathrm{d}t}s90 = |v_{36}| + |v_{49}| - |v_{23}| \tag{198}$$

### 10.9 Species species\_1

Name eIL17

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

### Initial assignment species\_1

This species takes part in three reactions (as a reactant in reaction\_3 and as a product in reaction\_2 and as a modifier in reaction\_3).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{-1} = v_{38} - v_{39} \tag{199}$$

### 10.10 Species species\_2

Name eIL10

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

### Initial assignment species\_2

This species takes part in five reactions (as a reactant in re39 and as a product in reaction\_1, reaction\_12 and as a modifier in re39, reaction\_12).

$$\frac{d}{dt} \text{species}_2 = |v_{37}| + |v_{48}| - |v_{32}| \tag{200}$$

### 10.11 Species species\_12

Name eIL18

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

Initial assignment species\_12

This species takes part in five reactions (as a reactant in re2 and as a product in reaction\_4, reaction\_6 and as a modifier in re2, reaction\_6).

$$\frac{d}{dt} \text{species}_{-12} = |v_{40}| + |v_{42}| - |v_1| \tag{201}$$

### 10.12 Species species\_13

Name anti-IL4

Initial concentration  $1 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a modifier in re20, re20), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{-}13 = 0 \tag{202}$$

### 10.13 Species species\_14

Name anti-IFNg

Initial concentration  $1 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a modifier in re14, re14), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{-}14 = 0 \tag{203}$$

## 10.14 Species species\_15

Name pIL4

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

This species takes part in two reactions (as a reactant in re20 and as a modifier in re20), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{-}15 = 0 \tag{204}$$

### 10.15 Species species\_16

Name IL18\_pool

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

This species takes part in two reactions (as a reactant in reaction\_6 and as a modifier in reaction\_6), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{-}16 = 0 \tag{205}$$

### **10.16 Species** species\_17

Name IL12\_pool

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

This species takes part in two reactions (as a reactant in reaction\_7 and as a modifier in reaction\_7), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{17} = 0 \tag{206}$$

### 10.17 Species species\_18

Name IFNg\_pool

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

This species takes part in two reactions (as a reactant in reaction\_8 and as a modifier in reaction\_8), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{1}18 = 0 \tag{207}$$

#### **10.18 Species** species\_19

Name IL21\_pool

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

This species takes part in two reactions (as a reactant in reaction\_9 and as a modifier in reaction\_9), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{1}9 = 0 \tag{208}$$

### 10.19 Species species\_20

Name IL23\_pool

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

This species takes part in two reactions (as a reactant in reaction\_10 and as a modifier in reaction\_10), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{d}{dt} \text{species} 20 = 0 \tag{209}$$

### 10.20 Species species\_21

Name IL17\_pool

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

This species takes part in two reactions (as a reactant in reaction\_11 and as a modifier in reaction\_11), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{21} = 0 \tag{210}$$

## 10.21 Species species\_22

Name IL10\_pool

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

This species takes part in two reactions (as a reactant in reaction\_12 and as a modifier in reaction\_12), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{d}{dt} species_2 = 0 (211)$$

### 10.22 Species species\_23

Name IL6\_pool

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

This species takes part in two reactions (as a reactant in reaction\_13 and as a modifier in reaction\_13), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{d}{dt} species_2 23 = 0 {212}$$

### 10.23 Species species\_24

Name IL2\_pool

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

This species takes part in two reactions (as a reactant in reaction\_14 and as a modifier in reaction\_14), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}.24 = 0 \tag{213}$$

### 10.24 Species species\_25

Name TGFb\_pool

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

This species takes part in two reactions (as a reactant in reaction\_15 and as a modifier in reaction\_15), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{25} = 0 \tag{214}$$

## 10.25 Species species\_26

Name IL4\_pool

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

This species takes part in two reactions (as a reactant in reaction\_16 and as a modifier in reaction\_16), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{d}{dt} \text{species}_2 = 0 \tag{215}$$

## **10.26 Species** species\_27

Name pIL10

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

This species takes part in two reactions (as a reactant in reaction\_17 and as a modifier in reaction\_17), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{d}{dt} \text{species}_2 = 0 \tag{216}$$

### **10.27 Species** s48

Name IL6-IL6R

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

This species takes part in eight reactions (as a product in re29 and as a modifier in re27, re27, re29, re30, re30, re36, re36).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}48 = v_{23} \tag{217}$$

## **10.28 Species** s47

Name IL6R

Initial concentration  $0.9999999518 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in re29 and as a modifier in re29).

$$\frac{d}{dt}s47 = -v_{23} \tag{218}$$

### **10.29 Species** s46

Name IL6

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

This species takes part in two reactions (as a reactant in re46 and as a modifier in re46).

$$\frac{d}{dt}s46 = -v_{36} \tag{219}$$

### **10.30 Species** s45

Name TGFb-TGFbR

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

This species takes part in ten reactions (as a product in re28 and as a modifier in re15, re15, re23, re23, re27, re27, re28, re31, re31).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}45 = |v_{22}|\tag{220}$$

## **10.31 Species** s44

Name TGFbR

**Initial concentration**  $0.999999951844375 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a reactant in re28 and as a modifier in re28).

$$\frac{d}{dt}s44 = -v_{22} \tag{221}$$

## **10.32 Species** s43

Name TGFb

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

This species takes part in two reactions (as a reactant in re44 and as a modifier in re44).

$$\frac{d}{dt}s43 = -v_{34} \tag{222}$$

#### **10.33 Species** s38

Name IL2-IL2R

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

This species takes part in four reactions (as a product in re25 and as a modifier in re25, re26, re26).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}38 = v_{19} \tag{223}$$

### **10.34 Species** s37

Name IL2R

Initial concentration  $0.999999951844375 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in re25 and as a modifier in re25).

$$\frac{d}{dt}s37 = -v_{19} \tag{224}$$

### **10.35 Species** s36

Name IL2

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

This species takes part in two reactions (as a reactant in re45 and as a modifier in re45).

$$\frac{d}{dt}s36 = -v_{35} \tag{225}$$

### **10.36 Species** s33

Name IL4-IL4R

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

This species takes part in six reactions (as a product in re19 and as a modifier in re19, re24, re24, re42, re42).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}33 = v_{15} \tag{226}$$

### **10.37 Species** s32

Name IL4R

Initial concentration  $0.999999951844375 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a reactant in re19 and as a modifier in re19).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}32 = -v_{15} \tag{227}$$

## **10.38 Species** s31

Name IL4

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

This species takes part in five reactions (as a reactant in re18 and as a product in re20 and as a modifier in re18, re20, re20).

$$\frac{\mathrm{d}}{\mathrm{d}t}s31 = |v_{16}| - |v_{14}| \tag{228}$$

### **10.39 Species** s25

Name IFNg-IFNgR

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

This species takes part in ten reactions (as a product in re11 and as a modifier in re11, re13, re13, re16, re16, re24, re24, re42, re42).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}25 = v_7 \tag{229}$$

### **10.40 Species** s24

Name IFNgR

**Initial concentration**  $0.999999951844375 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a reactant in re11 and as a modifier in re11).

$$\frac{\mathrm{d}}{\mathrm{d}t}s24 = -v_7\tag{230}$$

## **10.41 Species** s14

Name IL12-IL12R

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

This species takes part in four reactions (as a product in re8 and as a modifier in re8, re10, re10).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}14 = v_4 \tag{231}$$

### **10.42 Species** s13

Name IL12R

Initial concentration  $0.99999951844375 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in re8 and as a modifier in re8).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}13 = -v_4\tag{232}$$

### **10.43 Species** s12

Name IL12

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

This species takes part in two reactions (as a reactant in re6 and as a modifier in re6).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}12 = -v_3\tag{233}$$

### **10.44 Species** s3

Name IL18-IL18R

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

This species takes part in four reactions (as a product in re2 and as a modifier in re2, re3, re3).

$$\frac{\mathrm{d}}{\mathrm{d}t}s3 = v_1 \tag{234}$$

### **10.45 Species** s2

Name IL18R

Initial concentration  $0.999999951844375 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a reactant in re2 and as a modifier in re2).

$$\frac{\mathrm{d}}{\mathrm{d}t}s2 = -v_1\tag{235}$$

## 10.46 Species s1

Name IL18

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

This species takes part in two reactions (as a reactant in reaction\_4 and as a modifier in reaction\_4).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}1 = -v_{40} \tag{236}$$

## **10.47 Species** s52

Name IL21

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

This species takes part in four reactions (as a reactant in re33 and as a product in re37 and as a modifier in re33, re37).

$$\frac{\mathrm{d}}{\mathrm{d}t}s52 = |v_{31}| - |v_{27}| \tag{237}$$

### **10.48 Species** s54

Name IL21-IL21R

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

This species takes part in ten reactions (as a product in re32 and as a modifier in re10, re10, re13, re13, re30, re30, re32, reaction\_17, reaction\_17).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}54 = v_{26} \tag{238}$$

## **10.49 Species** s53

Name IL21R

**Initial concentration**  $0.99999951844375 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a reactant in re32 and as a modifier in re32).

$$\frac{d}{dt}s53 = -v_{26} \tag{239}$$

### **10.50 Species** s58

Name IL23R

Initial concentration  $0.99999951844375 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in re35 and as a modifier in re35).

$$\frac{d}{dt}s58 = -v_{29} \tag{240}$$

## **10.51 Species** s59

Name IL23-IL23R

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

This species takes part in ten reactions (as a product in re35 and as a modifier in re10, re10, re30, re30, re35, reaction\_5, reaction\_5, reaction\_17, reaction\_17).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}59 = |v_{29}|\tag{241}$$

## **10.52 Species** s62

Name IL10R

Initial concentration  $0.999999951844375 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a reactant in re39 and as a modifier in re39).

$$\frac{d}{dt}s62 = -v_{32} \tag{242}$$

### **10.53 Species** s63

Name IL10-IL10R

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

This species takes part in four reactions (as a product in re39 and as a modifier in re30, re30, re39).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}63 = v_{32} \tag{243}$$

### **10.54 Species** s65

Name IRAK1

Initial concentration  $0.5 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in re3 and as a modifier in re3), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}65 = 0\tag{244}$$

## **10.55 Species** s10

Name IRAK1-P

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

This species takes part in four reactions (as a product in re3 and as a modifier in re3, re9, re9).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}10 = v_2 \tag{245}$$

### **10.56 Species** s20

Name p50/p65 dimer

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

This species takes part in four reactions (as a product in re9 and as a modifier in re9, re14, re14).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}20 = v_5 \tag{246}$$

### **10.57 Species** s67

Name STAT4

Initial concentration  $0.999999951844375 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a reactant in re10 and as a modifier in re10), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}67 = 0\tag{247}$$

### **10.58 Species** s21

Name STAT4-P

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

This species takes part in eight reactions (as a product in re10 and as a modifier in re10, re14, re14, re15, re15, reaction\_5, reaction\_5).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}21 = v_6 \tag{248}$$

### **10.59 Species** s68

Name IFNg

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

This species takes part in four reactions (as a reactant in re12 and as a product in re14 and as a modifier in re12, re14).

$$\frac{d}{dt}s68 = |v_{10}| - |v_8| \tag{249}$$

### **10.60 Species** s69

Name JAK1

Initial concentration  $0.5 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in re16 and as a modifier in re16), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}69 = 0\tag{250}$$

### **10.61 Species** s28

Name JAK1-P

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

This species takes part in four reactions (as a product in re16 and as a modifier in re13, re13, re16).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}28 = v_{12} \tag{251}$$

## **10.62 Species** s70

Name STAT1

Initial concentration  $0.999999951844375 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in re13 and as a modifier in re13), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}70 = 0\tag{252}$$

### **10.63 Species** s26

Name STAT1-P

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

This species takes part in ten reactions (as a product in re13 and as a modifier in re13, re15, re15, re17, re17, re20, re20, re27, re27).

$$\frac{\mathrm{d}}{\mathrm{d}t}s26 = v_9 \tag{253}$$

### **10.64 Species** s57

Name p40/p19 dimer

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

This species takes part in six reactions (as a reactant in re35 and as a product in re34, re36 and as a modifier in re34, re35, re36).

$$\frac{\mathrm{d}}{\mathrm{d}t}s57 = |v_{28}| + |v_{30}| - |v_{29}| \tag{254}$$

### **10.65 Species** s73

Name IL17

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

This species takes part in six reactions (as a reactant in reaction\_2 and as a product in reaction\_5, reaction\_11 and as a modifier in reaction\_2, reaction\_5, reaction\_11).

$$\frac{\mathrm{d}}{\mathrm{d}t} s73 = |v_{41}| + |v_{47}| - |v_{38}| \tag{255}$$

### **10.66 Species** s74

Name IL10

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

This species takes part in four reactions (as a reactant in reaction\_1 and as a product in reaction\_17 and as a modifier in reaction\_1, reaction\_17).

$$\frac{\mathrm{d}}{\mathrm{d}t}s74 = |v_{53}| - |v_{37}| \tag{256}$$

### **10.67 Species** s75

Name RORgt

**Initial concentration**  $0.99999951844375 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a reactant in re31 and as a modifier in re31), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}75 = 0\tag{257}$$

### **10.68 Species** s50

Name RORgt-ligand

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

This species takes part in four reactions (as a product in re31 and as a modifier in re31, reaction\_5, reaction\_5).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}50 = v_{25} \tag{258}$$

### **10.69 Species** s49

Name STAT3-P

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

This species takes part in 14 reactions (as a product in re30 and as a modifier in re14, re14, re15, re15, re23, re23, re27, re27, re30, re31, re31, re37, re37).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}49 = v_{24} \tag{259}$$

### **10.70 Species** s76

Name STAT3

Initial concentration  $0.999999951844375 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in re30 and as a modifier in re30), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}76 = 0\tag{260}$$

## **10.71 Species** s39

Name STAT5-P

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

This species takes part in ten reactions (as a product in re26 and as a modifier in re20, re20, re26, re27, re32, re32, reaction\_17, reaction\_17).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}39 = v_{20} \tag{261}$$

### **10.72 Species** s77

Name STAT5

Initial concentration  $0.999999951844375 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a reactant in re26 and as a modifier in re26), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}77 = 0\tag{262}$$

### **10.73 Species** s78

Name FOXP3

Initial concentration  $0.999999951844375 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a reactant in re27 and as a modifier in re27), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}78 = 0\tag{263}$$

### **10.74 Species** s79

Name SOCS1

Initial concentration  $0.999999951844375 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a reactant in re17 and as a modifier in re17), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}79 = 0\tag{264}$$

### **10.75 Species** s29

Name SOCS1-JAKs

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

This species takes part in eight reactions (as a product in re17 and as a modifier in re11, re11, re16, re16, re17, re19, re19).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}29 = v_{13} \tag{265}$$

### **10.76 Species** s27

Name Tbet-P

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

This species takes part in eleven reactions (as a product in re15 and as a modifier in re14, re14, re15, re15, re17, re17, re23, re23, re35, re35).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}27 = |v_{11}|\tag{266}$$

### **10.77 Species** s80

Name Tbet

Initial concentration  $0.999999951844375 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a reactant in re15 and as a modifier in re15), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}80 = 0\tag{267}$$

### **10.78 Species** s81

Name GATA3

Initial concentration  $0.999999951844375 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a reactant in re23 and as a modifier in re23), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}81 = 0\tag{268}$$

## **10.79 Species** s35

Name GATA3-P

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

This species takes part in eight reactions (as a product in re23 and as a modifier in re10, re10, re20, re20, re23, reaction\_17, reaction\_17).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}35 = v_{17} \tag{269}$$

### **10.80 Species** s34

Name STAT6-P

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

This species takes part in ten reactions (as a product in re24 and as a modifier in re2, re2, re8, re8, re23, re23, re24, re27, re27).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}34 = v_{18} \tag{270}$$

### **10.81 Species** s82

Name STAT6

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

This species takes part in two reactions (as a reactant in re24 and as a modifier in re24), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}82 = 0\tag{271}$$

#### **10.82 Species** s85

Name PPARg

Initial concentration  $0.08 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in re42 and as a modifier in re42).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}85 = -v_{33} \tag{272}$$

## **10.83 Species** s83

Name L-PPARg

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

This species takes part in 26 reactions (as a product in re42 and as a modifier in re8, re9, re9, re13, re14, re14, re16, re16, re23, re23, re27, re27, re30, re30, re31, re31, re36, re36, re42, reaction\_5, reaction\_5, reaction\_17, reaction\_17).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}83 = v_{33} \tag{273}$$

#### **10.84 Species** s86

Name Ligand

Initial concentration  $0.9999999518 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in re42 and as a modifier in re42), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}s86 = 0\tag{274}$$

### **10.85 Species** s40

Name acetylated FOXP3

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

This species takes part in six reactions (as a product in re27 and as a modifier in re9, re9, re27, re31, re31).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{s}40 = v_{21} \tag{275}$$

## 10.86 Species species\_8

Name p19

Initial concentration  $0.9999999518 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a reactant in re36 and as a modifier in re36), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{-8} = 0 \tag{276}$$

### 10.87 Species species\_9

Name p40

Initial concentration  $0.9999999518 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in re36 and as a modifier in re36), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{9} = 0 \tag{277}$$

## 10.88 Species species\_10

Name p50

Initial concentration  $0.5 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in re9 and as a modifier in re9), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{-}10 = 0 \tag{278}$$

### 10.89 Species species\_11

Name p65

Initial concentration  $0.5 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a reactant in re9 and as a modifier in re9), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{-}11 = 0 \tag{279}$$

### 10.90 Species species\_4

Name IL17R

Initial concentration  $0.9999999518 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in reaction\_3 and as a modifier in reaction\_3).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{4} = -\nu_{39} \tag{280}$$

## 10.91 Species species\_3

Name IL17-IL17R

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

This species takes part in four reactions (as a product in reaction\_3 and as a modifier in re32, re32, reaction\_3).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{.3} = v_{39} \tag{281}$$

### 10.92 Species species\_5

Name pIFNg

Initial concentration  $0.9999999518 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in re14 and as a modifier in re14), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{d}{dt} species\_5 = 0 \tag{282}$$

## 10.93 Species species\_6

Name pIL21

**Initial concentration**  $0.99999951844375 \text{ mol} \cdot 1^{-1}$ 

This species takes part in two reactions (as a reactant in re37 and as a modifier in re37), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{-}6 = 0 \tag{283}$$

### **10.94 Species** species\_7

Name pIL17

Initial concentration  $0.999999951844375 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in reaction\_5 and as a modifier in reaction\_5), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{species}_{.7} = 0 \tag{284}$$

# A Glossary of Systems Biology Ontology Terms

**SBO:0000290 physical compartment:** Specific location of space, that can be bounded or not. A physical compartment can have 1, 2 or 3 dimensions

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