# **SBML Model Report**

# Model name: "Kallenberger2014 - CD95L induced apoptosis initiated by caspase-8, CD95 HeLa cells (cis/trans variant)"



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# 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 Stefan Kallenberger<sup>2</sup> at March thirteenth 2014 at 9:49 p. m. and last time modified at February 25<sup>th</sup> 2015 at 12:38 a. m. Table 1 shows an overview of the quantities of all components of this model.

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

Element	Quantity	Element	Quantity
compartment types	0	compartments	1
species types	0	species	18
events	0	constraints	0
reactions	13	function definitions	0
global parameters	12	unit definitions	0
rules	1	initial assignments	0

#### **Model Notes**

Kallenberger2014 - CD95L induced apoptosis initiated by caspase-8, CD95 HeLa cells (cis/trans variant)

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The paper describes a new approach that combines single cell and population data in the same model. The model consists of a large number of single cell models, which are fitted to single cell data. Simultaneously, ensemble averages are fitted to population data. It is assumed that the kinetics in each cell can be described with the same kinetic parameters. Therefore, cell-to-cell variability is explained by variable initial protein concentrations.

There are four variants of the model (with [CD95L]=500ng/ml = 16.6nM), i) cistrans (in CD95-HeLa cells) [ MODEL1403050000 ], ii) cistrans (in wild-type HeLa cells) [ MODEL1403050001 ], iii) cistrans-cistrans (in CD95-HeLa cells) [ MODEL1403050002 ], and iv) cistrans-cistrans (in wild-type HeLa cells) [ MODEL1403050003 ].

These model contain the equations for one "average cell, with median initial concentrations for CD95, FADD, p55, BID, PrNES\_mCherry and PrER\_mGFP. By integrating the model, it should be possible to obtain trajectories for PrER\_mGFP, PrNES\_mCherry, p43 and p18 similar as in Figure 4A (CD95-HeLa cells) and Figure 4B (wild-type HeLa cells).

This model is described in the article:Intra- and Interdimeric Caspase-8 Self-Cleavage Controls Strength and Timing of CD95-Induced ApoptosisStefan M. Kallenberger, Jol Beaudouin, Juliane Claus, Carmen Fischer, Peter K. Sorger, Stefan Legewie, and Roland Eils11 March 2014: Vol. 7, Issue 316, p. ra23

#### Abstract:

Apoptosis in response to the ligand CD95L (also known as Fas ligand) is initiated by caspase-8, which is activated by dimerization and self-cleavage at death-inducing signaling complexes (DISCs). Previous work indicated that the degree of substrate cleavage by caspase-8 determines whether a cell dies or survives in response to a death stimulus. To determine how a death ligand stimulus is effectively translated into caspase-8 activity, we assessed this activity over time in single cells with compartmentalized probes that are cleaved by caspase-8 and used multiscale modeling to simultaneously describe single-cell and population data with an ensemble of singlecell models. We derived and experimentally validated a minimal model in which cleavage of caspase-8 in the enzymatic domain occurs in an interdimeric manner through interaction between DISCs, whereas prodomain cleavage sites are cleaved in an intradimeric manner within DISCs. Modeling indicated that sustained membrane-bound caspase-8 activity is followed by transient cytosolic activity, which can be interpreted as a molecular timer mechanism reflected by a limited lifetime of active caspase-8. The activation of caspase-8 by combined intra- and interdimeric cleavage ensures weak signaling at low concentrations of CD95L and strongly accelerated activation at higher ligand concentrations, thereby contributing to precise control of apoptosis.

This model is hosted on BioModels Database and identifiedby: BIOMD0000000523.

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

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# 2 Unit Definitions

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

#### 2.1 Unit substance

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

**Definition** mol

#### 2.2 Unit volume

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

**Definition** 1

#### 2.3 Unit area

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

**Definition**  $m^2$ 

#### 2.4 Unit length

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

**Definition** m

#### 2.5 Unit time

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

**Definition** s

# 3 Compartment

This model contains one compartment.

Table 2: Properties of all compartments.

Id	Name	SBO	Spatial Dimensions	Size	Unit	Constant	Outside
cell	cell	0000290	3	1	litre		

# 3.1 Compartment cell

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

Name cell

SBO:0000290 physical compartment

# 4 Species

This model contains 18 species. Section 8 provides further details and the derived rates of change of each species.

Table 3: Properties of each species.

Id	Name	Compartment	Derived Unit	Constant	Boundary Condi- tion
CD95	CD95	cell	$\text{mol} \cdot 1^{-1}$		
FADD	FADD	cell	$\mathrm{mol}\cdot\mathrm{l}^{-1}$		$\Box$
p55free	p55free	cell	$\operatorname{mol} \cdot 1^{-1}$		$\Box$
Bid	Bid	cell	$\operatorname{mol} \cdot 1^{-1}$		$\Box$
PrNES_mCherry	PrNES_mCherry	cell	$\operatorname{mol} \cdot 1^{-1}$		
PrER_mGFP	PrER_mGFP	cell	$\operatorname{mol} \cdot 1^{-1}$		
DISC	DISC	cell	$\mathrm{mol}\cdot\mathrm{l}^{-1}$		
DISCp55	DISCp55	cell	$\mathrm{mol}\cdot\mathrm{l}^{-1}$		
p30	p30	cell	$\mathrm{mol}\cdot\mathrm{l}^{-1}$		$\Box$
p43	p43	cell	$\operatorname{mol} \cdot 1^{-1}$		
p18	p18	cell	$\operatorname{mol} \cdot 1^{-1}$		
p18inactive	p18inactive	cell	$\operatorname{mol} \cdot 1^{-1}$		
tBid	tBid	cell	$\operatorname{mol} \cdot 1^{-1}$		
PrNES	PrNES	cell	$\mathrm{mol}\cdot\mathrm{l}^{-1}$		
mCherry	mCherry	cell	$\mathrm{mol}\cdot\mathrm{l}^{-1}$		
PrER	PrER	cell	$\operatorname{mol} \cdot 1^{-1}$		
mGFP	mGFP	cell	$\text{mol} \cdot l^{-1}$		
CD95L	CD95L	cell	$\operatorname{mol} \cdot 1^{-1}$		

# **5 Parameters**

This model contains twelve global parameters.

Table 4: Properties of each parameter.

Id	Name	SBO	Value	Unit	Constant
kon_FADD		8.1	1711012144556 - 1	$10^{-4}$	$\overline{Z}$
$koff\_FADD$			0.006		$   \overline{\mathscr{L}} $
kDISC		4.9	1828591049766 - 1	$10^{-4}$	$   \overline{\mathscr{L}} $
kD216			0.011		$\overline{\mathbf{Z}}$
kD374trans-		4.4	6994772958953 - 1	$10^{-4}$	$\overline{\mathbf{Z}}$
_p55					
kD374trans-			0.003		
_p43					
kdiss_p18			0.095		
kBid		5	2867403363568 - 1	$10^{-4}$	$\overline{\mathbf{Z}}$
kD374probe			0.002		$\overline{\mathbf{Z}}$
KDR			8.985		$\overline{\mathbf{Z}}$
KDL			15.422		$\overline{\mathbf{Z}}$
CD95act			0.000		

# 6 Rule

This is an overview of one rule.

# 6.1 Rule CD95act

Rule CD95act is an assignment rule for parameter CD95act:

$$\begin{split} & \text{CD95act} \\ &= \frac{[\text{CD95}]^3 \cdot \text{KDL}^2 \cdot [\text{CD95L}]}{\left([\text{CD95L}] + \text{KDL}\right) \cdot \left([\text{CD95}]^2 \cdot \text{KDL}^2 + \text{KDR} \cdot [\text{CD95L}]^2 + 2 \cdot \text{KDR} \cdot \text{KDL} \cdot [\text{CD95L}] + \text{KDR} \cdot \text{KDL}^2\right)} \end{split}$$

# 7 Reactions

This model contains 13 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

N⁰	Id Name	Reaction Equation SBO
1	reaction_1	$FADD \xrightarrow{FADD} DISC$
2	reaction_2	$\overline{\mathrm{DISC}} \xrightarrow{\overline{\mathrm{DISC}}} \mathrm{FADD}$
3	reaction_3	p55free + DISC $\xrightarrow{\text{p55free}}$ DISCp55
4	reaction_4	DISCp55 $\xrightarrow{\text{DISCp55}}$ p30
5	reaction_5	$p43 \xrightarrow{p43} p18 + DISC$
6	reaction_6	DISCp55 $\xrightarrow{\text{DISCp55}}$ , p30, DISCp55, p30 p43
7	reaction_7	DISCp55 $\xrightarrow{\text{p43}}$ DISCp55, p43 $\xrightarrow{\text{p43}}$ p43
8	reaction_8	$p30 \xrightarrow{DISCp55, p30, p30, DISCp55} p18 + DISC$
9	reaction_9	$p30 \xrightarrow{p43, p30, p43} p18 + DISC$
10	reaction_10	$p18 \xrightarrow{p18} p18$ inactive
11		$\text{Bid} \xrightarrow{\text{p43, p18, Bid, p43, p18}} \text{tBid}$
11	reaction_11	
12	reaction_12	PrNES_mCherry $\xrightarrow{p43, p18, PrNES\_mCherry, p43, p18} PrNES +$
		mCherry
13	reaction_13	$PrER\_mGFP \xrightarrow{p18, PrER\_mGFP, p18} PrER + mGFP$

#### 7.1 Reaction reaction\_1

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

# **Reaction equation**

$$FADD \xrightarrow{FADD} DISC \tag{2}$$

#### Reactant

Table 6: Properties of each reactant.

Id	Name	SBO
FADD	FADD	

#### **Modifier**

Table 7: Properties of each modifier.

Id	Name	SBO
FADD	FADD	

#### **Product**

Table 8: Properties of each product.

Id	Name	SBO
DISC	DISC	

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_1 = \text{kon\_FADD} \cdot \text{CD95act} \cdot [\text{FADD}] \cdot \text{vol} (\text{cell})$$
 (3)

#### 7.2 Reaction reaction\_2

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

# **Reaction equation**

$$DISC \xrightarrow{DISC} FADD \tag{4}$$

#### Reactant

Table 9: Properties of each reactant.

Id	Name	SBO
DISC	DISC	

#### **Modifier**

Table 10: Properties of each modifier.

Id	Name	SBO
DISC	DISC	

#### **Product**

Table 11: Properties of each product.

Id	Name	SBO
FADD	FADD	

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_2 = \text{koff\_FADD} \cdot [\text{DISC}] \cdot \text{vol} (\text{cell})$$
 (5)

# 7.3 Reaction reaction\_3

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

# **Reaction equation**

$$p55free + DISC \xrightarrow{p55free, DISC} DISCp55$$
 (6)

#### **Reactants**

Table 12: Properties of each reactant.

Id	Name	SBO
p55free	p55free	

Id	Name	SBO
DISC	DISC	

#### **Modifiers**

Table 13: Properties of each modifier.

Id	Name	SBO
p55free DISC	p55free DISC	

#### **Product**

Table 14: Properties of each product.

Id	Name	SBO
DISCp55	DISCp55	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_3 = \text{kDISC} \cdot [\text{p55free}] \cdot [\text{DISC}] \cdot \text{vol} (\text{cell})$$
 (7)

# 7.4 Reaction reaction\_4

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

# **Reaction equation**

$$DISCp55 \xrightarrow{DISCp55} p30 \tag{8}$$

#### Reactant

Table 15: Properties of each reactant.

Id	Name	SBO
DISCp55	DISCp55	

# Modifier

Table 16: Properties of each modifier.

Id	Name	SBO
DISCp55	DISCp55	

#### **Product**

Table 17: Properties of each product.

Id	Name	SBO
p30	p30	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_4 = \text{kD216} \cdot [\text{DISCp55}] \cdot \text{vol}(\text{cell}) \tag{9}$$

#### 7.5 Reaction reaction\_5

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

# **Reaction equation**

$$p43 \xrightarrow{p43} p18 + DISC \tag{10}$$

#### Reactant

Table 18: Properties of each reactant.

Id	Name	SBO
p43	p43	

# **Modifier**

Table 19: Properties of each modifier.

Id	Name	SBO
p43	p43	

#### **Products**

Table 20: Properties of each product.

Id	Name	SBO
p18 DISC	p18 DISC	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_5 = \text{kD216} \cdot [\text{p43}] \cdot \text{vol}(\text{cell}) \tag{11}$$

# **7.6 Reaction** reaction\_6

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

# **Reaction equation**

DISCp55 
$$\xrightarrow{\text{DISCp55}}$$
, p30, DISCp55, p30 p43 (12)

#### Reactant

Table 21: Properties of each reactant.

Id	Name	SBO
DISCp55	DISCp55	

#### **Modifiers**

Table 22: Properties of each modifier.

Id	Name	SBO
DISCp55	DISCp55	
p30	p30	
DISCp55	DISCp55	
p30	p30	

# **Product**

Table 23: Properties of each product.

Id	Name	SBO
p43	p43	

**Derived unit** contains undeclared units

$$v_6 = \text{kD374trans\_p55} \cdot [\text{DISCp55}] \cdot ([\text{DISCp55}] + [\text{p30}]) \cdot \text{vol}(\text{cell})$$
(13)

#### 7.7 Reaction reaction\_7

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

# **Reaction equation**

$$DISCp55 \xrightarrow{p43, DISCp55, p43} p43$$
 (14)

#### Reactant

Table 24: Properties of each reactant.

Id	Name	SBO
DISCp55	DISCp55	

# **Modifiers**

Table 25: Properties of each modifier.

Id	Name	SBO
p43	p43	
DISCp55	DISCp55	
p43	p43	

#### **Product**

Table 26: Properties of each product.

Id	Name	SBO
p43	p43	

**Derived unit** contains undeclared units

$$v_7 = \text{kD374trans\_p43} \cdot [\text{DISCp55}] \cdot [\text{p43}] \cdot \text{vol} (\text{cell})$$
 (15)

#### 7.8 Reaction reaction\_8

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

# **Reaction equation**

$$p30 \xrightarrow{DISCp55, p30, p30, DISCp55} p18 + DISC$$
 (16)

#### Reactant

Table 27: Properties of each reactant.

Id	Name	SBO
p30	p30	

#### **Modifiers**

Table 28: Properties of each modifier.

Name	SBO
DISCp55	
p30	
p30	
DISCp55	
	DISCp55 p30 p30

#### **Products**

Table 29: Properties of each product.

Id	Name	SBO
p18 DISC	p18 DISC	

**Derived unit** contains undeclared units

$$v_8 = \text{kD374trans\_p55} \cdot [\text{p30}] \cdot ([\text{DISCp55}] + [\text{p30}]) \cdot \text{vol}(\text{cell})$$
(17)

#### 7.9 Reaction reaction\_9

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

# **Reaction equation**

$$p30 \xrightarrow{p43, p30, p43} p18 + DISC$$
 (18)

#### Reactant

Table 30: Properties of each reactant.

Id	Name	SBO
p30	p30	

#### **Modifiers**

Table 31: Properties of each modifier.

Id	Name	SBO
p43	p43	
p30	p30	
p43	p43	

#### **Products**

Table 32: Properties of each product.

Id	Name	SBO
p18 DISC	p18 DISC	

**Derived unit** contains undeclared units

$$v_9 = \text{kD374trans\_p43} \cdot [\text{p30}] \cdot [\text{p43}] \cdot \text{vol(cell)}$$
(19)

#### 7.10 Reaction reaction\_10

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

# **Reaction equation**

$$p18 \xrightarrow{p18} p18 inactive \tag{20}$$

#### Reactant

Table 33: Properties of each reactant.

Id	Name	SBO
p18	p18	

#### **Modifier**

Table 34: Properties of each modifier.

Id	Name	SBO
p18	p18	

#### **Product**

Table 35: Properties of each product.

1	1	
Id	Name	SBO
p18inactive	p18inactive	

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{10} = \text{kdiss\_p18} \cdot [\text{p18}] \cdot \text{vol}(\text{cell})$$
 (21)

# 7.11 Reaction reaction\_11

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

# **Reaction equation**

$$Bid \xrightarrow{p43, p18, Bid, p43, p18} tBid$$
 (22)

#### Reactant

Table 36: Properties of each reactant.

Id	Name	SBO
Bid	Bid	

#### **Modifiers**

Table 37: Properties of each modifier.

Id	Name	SBO
p43	p43	
p18	p18	
Bid	Bid	
p43	p43	
p18	p18	

#### **Product**

Table 38: Properties of each product.

Id	Name	SBO
tBid	tBid	

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{11} = \text{kBid} \cdot [\text{Bid}] \cdot ([\text{p43}] + [\text{p18}]) \cdot \text{vol}(\text{cell})$$
(23)

#### 7.12 Reaction reaction\_12

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

# **Reaction equation**

$$PrNES\_mCherry \xrightarrow{p43, p18, PrNES\_mCherry, p43, p18} PrNES + mCherry$$
 (24)

#### Reactant

Table 39: Properties of each reactant.

Table 37. I roperties of each reactain.		
Id	Name	SBO
PrNES_mCherry	PrNES_mCherry	<u>.</u>

#### **Modifiers**

Table 40: Properties of each modifier.

Id	Name	SBO
p43	p43	
p18	p18	
PrNES_mCherry	PrNES_mCherry	
p43	p43	
p18	p18	

#### **Products**

Table 41: Properties of each product.

Id	Name	SBO
PrNES	PrNES	
mCherry	mCherry	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{12} = \text{kD374probe} \cdot [\text{PrNES\_mCherry}] \cdot ([\text{p43}] + [\text{p18}]) \cdot \text{vol} (\text{cell})$$
 (25)

#### 7.13 Reaction reaction\_13

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

#### **Reaction equation**

$$PrER\_mGFP \xrightarrow{p18, PrER\_mGFP, p18} PrER + mGFP$$
 (26)

#### Reactant

Table 42: Properties of each reactant.

Id	Name	SBO
PrER_mGFP	PrER_mGFP	

#### **Modifiers**

Table 43: Properties of each modifier.

Id	Name	SBO
p18 PrER_mGFP p18	p18 PrER_mGFP p18	

#### **Products**

Table 44: Properties of each product.

Id	Name	SBO
PrER	PrER	
mGFP	mGFP	

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{13} = \text{kD374probe} \cdot [\text{PrER\_mGFP}] \cdot [\text{p18}] \cdot \text{vol}(\text{cell})$$
 (27)

# 8 Derived Rate Equations

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

Identifiers for kinetic laws highlighted in gray cannot be verified to evaluate to units of SBML substance per time. As a result, some SBML interpreters may not be able to verify the consistency of the units on quantities in the model. Please check if

- parameters without an unit definition are involved or
- volume correction is necessary because the hasOnlySubstanceUnits flag may be set to false and spacialDimensions> 0 for certain species.

#### 8.1 Species CD95

Name CD95

Initial concentration  $116 \text{ mol} \cdot l^{-1}$ 

This species does not take part in any reactions. Its quantity does hence not change over time:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{CD95} = 0\tag{28}$$

#### 8.2 Species FADD

Name FADD

Initial concentration  $93 \text{ mol} \cdot l^{-1}$ 

This species takes part in three reactions (as a reactant in reaction\_1 and as a product in reaction\_2 and as a modifier in reaction\_1).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{FADD} = v_2 - v_1 \tag{29}$$

# 8.3 Species p55free

Name p55free

Initial concentration  $155 \text{ mol} \cdot l^{-1}$ 

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

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{p55free} = -v_3\tag{30}$$

# 8.4 Species Bid

Name Bid

Initial concentration  $236 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in reaction\_11 and as a modifier in reaction\_11).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Bid} = -v_{11} \tag{31}$$

#### 8.5 Species PrNES\_mCherry

Name PrNES\_mCherry

Initial concentration  $973 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in reaction\_12 and as a modifier in reaction\_12).

$$\frac{d}{dt} PrNES\_mCherry = -v_{12}$$
 (32)

#### 8.6 Species PrER\_mGFP

Name PrER\_mGFP

Initial concentration  $5178 \text{ mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in reaction\_13 and as a modifier in reaction\_13).

$$\frac{\mathrm{d}}{\mathrm{d}t} \text{PrER}_{-} \text{mGFP} = -v_{13} \tag{33}$$

#### 8.7 Species DISC

Name DISC

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

This species takes part in eight reactions (as a reactant in reaction\_2, reaction\_3 and as a product in reaction\_1, reaction\_5, reaction\_8, reaction\_9 and as a modifier in reaction\_2, reaction\_3).

$$\frac{d}{dt}DISC = |v_1| + |v_5| + |v_8| + |v_9| - |v_2| - |v_3|$$
(34)

#### 8.8 Species DISCp55

Name DISCp55

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

This species takes part in ten reactions (as a reactant in reaction\_4, reaction\_6, reaction\_7 and as a product in reaction\_3 and as a modifier in reaction\_4, reaction\_6, reaction\_6, reaction\_7, reaction\_8, reaction\_8).

$$\frac{d}{dt}DISCp55 = |v_3| - |v_4| - |v_6| - |v_7|$$
(35)

#### 8.9 Species p30

Name p30

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

This species takes part in eight reactions (as a reactant in reaction\_8, reaction\_9 and as a product in reaction\_4 and as a modifier in reaction\_6, reaction\_6, reaction\_8, reaction\_8, reaction\_9).

$$\frac{d}{dt}p30 = |v_4| - |v_8| - |v_9| \tag{36}$$

#### **8.10 Species** p43

Name p43

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

This species takes part in twelve reactions (as a reactant in reaction\_5 and as a product in reaction\_6, reaction\_7 and as a modifier in reaction\_5, reaction\_7, reaction\_7, reaction\_9, reaction\_9, reaction\_11, reaction\_11, reaction\_12, reaction\_12).

$$\frac{d}{dt}p43 = |v_6| + |v_7| - |v_5| \tag{37}$$

# **8.11 Species** p18

Name p18

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

This species takes part in eleven reactions (as a reactant in reaction\_10 and as a product in reaction\_5, reaction\_8, reaction\_9 and as a modifier in reaction\_10, reaction\_11, reaction\_11, reaction\_12, reaction\_12, reaction\_13, reaction\_13).

$$\frac{\mathrm{d}}{\mathrm{d}t}p18 = |v_5| + |v_8| + |v_9| - |v_{10}| \tag{38}$$

#### 8.12 Species p18inactive

Name p18inactive

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

This species takes part in one reaction (as a product in reaction\_10).

$$\frac{\mathrm{d}}{\mathrm{d}t} p18 \text{inactive} = v_{10} \tag{39}$$

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#### 8.13 Species tBid

Name tBid

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

This species takes part in one reaction (as a product in reaction\_11).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathbf{t}\mathbf{B}\mathbf{i}\mathbf{d} = v_{11} \tag{40}$$

# 8.14 Species PrNES

Name PrNES

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

This species takes part in one reaction (as a product in reaction\_12).

$$\frac{\mathrm{d}}{\mathrm{d}t} \text{PrNES} = |v_{12}| \tag{41}$$

#### 8.15 Species mCherry

Name mCherry

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

This species takes part in one reaction (as a product in reaction\_12).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{mCherry} = v_{12} \tag{42}$$

#### 8.16 Species PrER

Name PrER

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

This species takes part in one reaction (as a product in reaction\_13).

$$\frac{\mathrm{d}}{\mathrm{d}t} \text{PrER} = v_{13} \tag{43}$$

# 8.17 Species mGFP

Name mGFP

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

This species takes part in one reaction (as a product in reaction\_13).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{mGFP} = v_{13} \tag{44}$$

# 8.18 Species CD95L

Name CD95L

Initial concentration  $16.6 \text{ mol} \cdot l^{-1}$ 

This species does not take part in any reactions. Its quantity does hence not change over time:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{CD95L} = 0\tag{45}$$

# 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

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

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