# **SBML Model Report**

# Model name: "Tang2010\_PolyGlutamate"



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 Carole J Proctor<sup>2</sup> at September 27<sup>th</sup> 2010 at 12:21 a.m. and last time modified at June third 2014 at 9:03 p.m. Table 1 gives an overview of the quantities of all components of this model.

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

Element	Quantity	Element	Quantity
compartment types	0	compartments	1
species types	0	species	27
events	2	constraints	0
reactions	72	function definitions	0
global parameters	48	unit definitions	1
rules	1	initial assignments	0

#### **Model Notes**

This a model from the article:

### Experimental and computational analysis of polyglutamine-mediated cytotoxicity.

Tang MY, Proctor CJ, Woulfe J, Gray DA. <u>PLoS Comput Biol.</u>2010 Sep 23;6(9). 20885783, **Abstract:** 

Expanded polyglutamine (polyQ) proteins are known to be the causative agents of a number

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of human neurodegenerative diseases but the molecular basis of their cytoxicity is still poorly understood. PolyQ tracts may impede the activity of the proteasome, and evidence from single cell imaging suggests that the sequestration of polyQ into inclusion bodies can reduce the proteasomal burden and promote cell survival, at least in the short term. The presence of misfolded protein also leads to activation of stress kinases such as p38MAPK, which can be cytotoxic. The relationships of these systems are not well understood. We have used fluorescent reporter systems imaged in living cells, and stochastic computer modeling to explore the relationships of polyQ, p38MAPK activation, generation of reactive oxygen species (ROS), proteasome inhibition, and inclusion body formation. In cells expressing a polyQ protein inclusion, body formation was preceded by proteasome inhibition but cytotoxicity was greatly reduced by administration of a p38MAPK inhibitor. Computer simulations suggested that without the generation of ROS, the proteasome inhibition and activation of p38MAPK would have significantly reduced toxicity. Our data suggest a vicious cycle of stress kinase activation and proteasome inhibition that is ultimately lethal to cells. There was close agreement between experimental data and the predictions of a stochastic computer model, supporting a central role for proteasome inhibition and p38MAPK activation in inclusion body formation and ROS-mediated cell death.

This model originates from BioModels Database: A Database of Annotated Published Models (http://www.ebi.ac.uk/biomodels/). It is copyright (c) 2005-2010 The BioModels.net Team. For more information see the terms of use.

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

### 2 Unit Definitions

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

#### 2.1 Unit substance

**Definition** item

### 2.2 Unit volume

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

**Definition** 1

#### 2.3 Unit area

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

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

# 2.4 Unit length

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

**Definition** m

### 2.5 Unit time

Notes Second is the predefined SBML unit for time.

**Definition** s

# 3 Compartment

This model contains one compartment.

Table 2: Properties of all compartments.

Id	Name	SBO	Spatial Dimensions	Size	Unit	Constant	Outside
cytosol			3	1	litre	Z	

# 3.1 Compartment cytosol

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

# 4 Species

This model contains 27 species. The boundary condition of two of these species is set to true so that these species' amount cannot be changed by any reaction. Section 9 provides further details and the derived rates of change of each species.

Table 3: Properties of each species.

	Compartment	Derived Unit	Constant	Boundary Condi-
				tion
PolyQ	cytosol	item		
Proteasome	cytosol	item		
NatP	cytosol	item		$\Box$
MisP	cytosol	item		$\Box$
MisP_Proteasome	cytosol	item		
AggP1	cytosol	item		
AggP2	cytosol	item		$\Box$
AggP3	cytosol	item		$\Box$
AggP4	cytosol	item		
AggP5	cytosol	item	$\Box$	$\Box$
AggPolyQ1	cytosol	item		
AggPolyQ2	cytosol	item	$\Box$	$\Box$
AggPolyQ3	cytosol	item	$\Box$	$\Box$
AggPolyQ4	cytosol	item		
AggPolyQ5	cytosol	item		$\Box$
SeqAggP	cytosol	item		$\Box$
AggP_Proteasome	cytosol	item		$\Box$
mRFPu	cytosol	item		$\Box$
mRFPu_Proteasome	cytosol	item		$\Box$
PolyQ_Proteasome	cytosol	item		
ROS	cytosol	item		

Id	Name	Compartment	Derived Unit	Constant	Boundary Condi- tion
p38_P		cytosol	item	В	$\Box$
p38		cytosol	item		
Source		cytosol	item	$\square$	$\square$
Sink		cytosol	item	$\square$	$\square$
p38death		cytosol	item		
PIdeath		cytosol	item	$\Box$	$\Box$

# **5 Parameters**

This model contains 48 global parameters.

Table 4: Properties of each parameter.

Id	Name	SBO	Value	Unit	Constant
kaggPolyQ			$5 \cdot 10^{-8}$		$\overline{Z}$
kdisaggPol	yQ1		$5\cdot 10^{-7}$		$ \overline{\checkmark} $
kdisaggPol	yQ2		$4 \cdot 10^{-7}$		
kdisaggPol	y <b>Q</b> 3		$3 \cdot 10^{-7}$		
kdisaggPol	yQ4		$2 \cdot 10^{-7}$		
kdisaggPol	y <b>Q</b> 5		$10^{-7}$		
${\tt kseqPolyQ}$			$8 \cdot 10^{-7}$		
kinhprot			$5\cdot 10^{-9}$		
kaggMisP			$10^{-11}$		
kagg2MisP			$10^{-10}$		
kdisaggMis	P1		$5 \cdot 10^{-7}$		
kdisaggMis	P2		$4 \cdot 10^{-7}$		
kdisaggMis			$3 \cdot 10^{-7}$		
kdisaggMis	P4		$2 \cdot 10^{-7}$		$   \overline{\checkmark} $
kdisaggMis	P5		$10^{-7}$		
ksynmRFPu			0.138		
kbinmRFPu			$5 \cdot 10^{-7}$		
krelmRFPu			$10^{-8}$		
kdegmRFPu			0.005		
${\tt ksynPolyQ}$			0.007		
${\tt kbinPolyQ}$			$5 \cdot 10^{-8}$		
krelPolyQ			$10^{-9}$		
${\tt kdegPolyQ}$			0.003		
kgenROS			0.002		
kremROS			$2 \cdot 10^{-4}$		
kgenROSAgg	P		$5 \cdot 10^{-6}$		
kgenROSSeq.	AggP		$10^{-7}$		$\square$
kactp38			$5 \cdot 10^{-6}$		$\square$
kinactp38			0.002		
${\tt kseqMisP}$			$10^{-9}$		
kseqAggPPr	ot		$5 \cdot 10^{-7}$		
kseqPolyQP:	rot		$5 \cdot 10^{-7}$		$\checkmark$
kseqMisPPr	ot		$5 \cdot 10^{-7}$		$\checkmark$
kseqmRFPuP:	rot		$5 \cdot 10^{-7}$		
${\tt kseqmRFPu}$			$10^{-10}$		$\checkmark$
${\tt ksynNatP}$			2.400		
kmisfold			$2 \cdot 10^{-6}$		

Id	Name	SBO	Value	Unit	Constant
krefold			$8 \cdot 10^{-5}$		lacksquare
${\tt kbinMisPProt}$			$5 \cdot 10^{-8}$		$\overline{\mathbf{Z}}$
krelMisPProt			$10^{-8}$		
kdegMisP			0.010		$\overline{\mathbf{Z}}$
kgenROSp38			$7 \cdot 10^{-4}$		$\overline{\mathbf{Z}}$
kp38act			1.000		
kp38death			$9 \cdot 10^{-8}$		
kPIdeath			$2.5 \cdot 10^{-8}$		
kproteff			1.000		$\Box$
kalive			1.000		
oligomers	oligomers		0.000		

### 6 Rule

This is an overview of one rule.

### **6.1 Rule** oligomers

Rule oligomers is an assignment rule for parameter oligomers:

$$oligomers = AggPolyQ1 + AggPolyQ2 + AggPolyQ3 + AggPolyQ4 + AggPolyQ5 \quad (1) \\$$

**Derived unit** item

### 7 Events

This is an overview of two events. Each event is initiated whenever its trigger condition switches from false to true. A delay function postpones the effects of an event to a later time point. At the time of execution, an event can assign values to species, parameters or compartments if these are not set to constant.

### 7.1 Event PIcellDeath

### **Trigger condition**

$$PIdeath > 0 (2)$$

### **Assignment**

$$kalive = 0 (3)$$

# **7.2 Event** p38cellDeath

#### 

# 8 Reactions

This model contains 72 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

		Table 5. Overview of an reactions			
$N_{\bar{0}}$	Id Name	Reaction Equation SBO			
1	polyQSynthesis	$Source \longrightarrow PolyQ$			
2	polyqProteasomeBinding	$PolyQ + Proteasome \longrightarrow PolyQ_Proteasome$			
3	polyqProteasomeRelease	$PolyQ_Proteasome \longrightarrow PolyQ + Proteasome$			
4	PolyQDegradation	PolyQ_Proteasome → Proteasome			
5	mRFPuSynthesis	Source $\longrightarrow$ mRFPu			
6	mRFPuProteasomeBinding	$mRFPu+Proteasome \longrightarrow mRFPu\_Proteasome$			
7	mRFPuProteasomeRelease	$mRFPu\_Proteasome \longrightarrow mRFPu+Proteasome$			
8	mRFPuDegradation	$mRFPu\_Proteasome \longrightarrow Proteasome$			
9	PolyQAggregation1	$2 \operatorname{PolyQ} + \operatorname{ROS} \longrightarrow \operatorname{AggPolyQ1} + \operatorname{ROS}$			
10	PolyQAggregation2	$PolyQ + AggPolyQ1 + ROS \longrightarrow AggPolyQ2 + ROS$			
11	PolyQAggregation3	$PolyQ + AggPolyQ2 + ROS \longrightarrow AggPolyQ3 + ROS$			
12	PolyQAggregation4	$PolyQ + AggPolyQ3 + ROS \longrightarrow AggPolyQ4 + ROS$			
13	PolyQAggregation5	$PolyQ + AggPolyQ4 + ROS \longrightarrow AggPolyQ5 + ROS$			
14	PolyQDisaggregation5	tion5 $AggPolyQ5 \longrightarrow PolyQ + AggPolyQ4$			
15	PolyQDisaggregation4	$AggPolyQ4 \longrightarrow PolyQ + AggPolyQ3$			
16	PolyQDisaggregation3	$AggPolyQ3 \longrightarrow PolyQ + AggPolyQ2$			
17	PolyQDisaggregation2	$AggPolyQ2 \longrightarrow PolyQ + AggPolyQ1$			
18	PolyQDisaggregation1 $AggPolyQ1 \longrightarrow 2 PolyQ$				
19	$ \texttt{PolyQInclusionFormation} \qquad \qquad \texttt{PolyQ} + \texttt{AggPolyQ5} \longrightarrow \texttt{7}  \texttt{SeqAggP} $				
20	PolyQInclusionGrowth	$\operatorname{PolyQ} + \operatorname{SeqAggP} \longrightarrow 2\operatorname{SeqAggP}$			
21	ProteasomeInhibition1	$\verb roteasomeInhibition1  AggPolyQ1 + Proteasome \longrightarrow AggP\_Proteasome $			
22	ProteasomeInhibition2	$AggPolyQ2 + Proteasome \longrightarrow AggP\_Proteasome$			
23	ProteasomeInhibition3	$AggPolyQ3 + Proteasome \longrightarrow AggP\_Proteasome$			

N⁰	Id Name	Reaction Equation SBO		
24	ProteasomeInhibition4	AggPolyQ4 + Proteasome → AggP_Proteasome		
25	ProteasomeInhibition5	$AggPolyQ5 + Proteasome \longrightarrow AggP\_Proteasome$		
26	mRFPuProteasomeSequestering	mRFPu_Proteasome + SeqAggP $\longrightarrow$ 2 SeqAggP		
27	mRFPuSequestering	$mRFPu + SeqAggP \longrightarrow 2 SeqAggP$		
28	ROSgenerationBasal	$Source \longrightarrow ROS$		
29	ROSgenerationSmallAggPolyQ1	$AggPolyQ1 \longrightarrow AggPolyQ1 + ROS$		
30	ROSgenerationSmallAggPolyQ2	$AggPolyQ2 \longrightarrow AggPolyQ2 + ROS$		
31	ROSgenerationSmallAggPolyQ3	$AggPolyQ3 \longrightarrow AggPolyQ3 + ROS$		
32	ROSgenerationSmallAggPolyQ4	$AggPolyQ4 \longrightarrow AggPolyQ4 + ROS$		
33	ROSgenerationSmallAggPolyQ5	$AggPolyQ5 \longrightarrow AggPolyQ5 + ROS$		
34	ROSgenerationAggPProteasome	$AggP\_Proteasome \longrightarrow AggP\_Proteasome + ROS$		
35	ROSremoval	$ROS \longrightarrow Sink$		
36	p38activation	$ROS + p38 \longrightarrow ROS + p38 P$		
37	p38inactivation	$p38.P \longrightarrow p38$		
38	AggP-	$AggP$ _Proteasome + Seq $AggP \longrightarrow 2$ Seq $AggP$		
	_ProteasomeSequestering			
39	PolyQ-	PolyQ_Proteasome + SeqAggP $\longrightarrow$ 2 SeqAggP		
	_ProteasomeSequestering	, , , , , , , , , , , , , , , , , , , ,		
40	MisP-	$MisP\_Proteasome + SeqAggP \longrightarrow 2 SeqAggP$		
	_ProteasomeSequestering	, 60		
41	ProteinSynthesis	Source $\longrightarrow$ NatP		
42	Misfolding	$NatP + ROS \longrightarrow MisP + ROS$		
43	Refolding	MisP → NatP		
44	MisP-	MisP+Proteasome → MisP-Proteasome		
	_ProteasomeBinding			
45	MisP-	$MisP\_Proteasome \longrightarrow MisP+Proteasome$		
	_ProteasomeRelease			
46	MisP-	MisP_Proteasome → Proteasome		
	_Degradation			

Nº	Id Name	Reaction Equation	SBO
47	MisP-	$2 \operatorname{MisP} \longrightarrow \operatorname{AggP1}$	
	_Aggregation1		
48	MisP-	$MisP + AggP1 \longrightarrow AggP2$	
	_Aggregation2		
49	MisP-	$MisP + AggP2 \longrightarrow AggP3$	
	_Aggregation3		
50	MisP-	$MisP + AggP3 \longrightarrow AggP4$	
	_Aggregation4		
51	MisP-	$MisP + AggP4 \longrightarrow AggP5$	
	$\_$ Aggregation5		
52	MisP-	$AggP1 \longrightarrow 2 MisP$	
	_Disaggregation1		
53	MisP-	$AggP2 \longrightarrow MisP + AggP1$	
	_Disaggregation2		
54	MisP-	$AggP3 \longrightarrow MisP + AggP2$	
	_Disaggregation3		
55	MisP-	$AggP4 \longrightarrow MisP + AggP3$	
	_Disaggregation4	A D5 A5 D A D4	
56	MisP-	$AggP5 \longrightarrow MisP + AggP4$	
	_Disaggregation5	M. D. A. D. G. G. A. D.	
57	MisP-	$MisP + AggP5 \longrightarrow 7 SeqAggP$	
<b>5</b> 0	_InclusionFormation	M'-D + CAD 2CAD	
58 50	MisPInclusionGrowth	$MisP + SeqAggP \longrightarrow 2 SeqAggP$	
59 60	ProteasomeInhibitionAggP1	$AggP1 + Proteasome \longrightarrow AggP\_Proteasome$	
	ProteasomeInhibitionAggP2	$AggP2 + Proteasome \longrightarrow AggP_Proteasome$	
61	ProteasomeInhibitionAggP3	$AggP3 + Proteasome \longrightarrow AggP_Proteasome$	
62 63	ProteasomeInhibitionAggP4	$AggP4 + Proteasome \longrightarrow AggP_Proteasome$	
64	ProteasomeInhibitionAggP5	$AggP5 + Proteasome \longrightarrow AggP\_Proteasome$ $AggP1 \longrightarrow AggP1 + ROS$	
04	ROSgenerationSmallAggP1	$Aggr1 \longrightarrow Aggr1 + ROS$	

N⁰	Id Name	Reaction Equation SBO
65	ROSgenerationSmallAggP2	$AggP2 \longrightarrow AggP2 + ROS$
66	ROSgenerationSmallAggP3	$AggP3 \longrightarrow AggP3 + ROS$
67	ROSgenerationSmallAggP4	$AggP4 \longrightarrow AggP4 + ROS$
68	ROSgenerationSmallAggP5	$AggP5 \longrightarrow AggP5 + ROS$
69	p38_P_ROS-	$p38\_P \longrightarrow p38\_P + ROS$
	_Generation	
70	ROSgenerationSeqAggP	$SeqAggP \longrightarrow SeqAggP + ROS$
71	P38DeathPathway	$p38\_P \longrightarrow p38\_P + p38$ death
72	PIDeathPathway	$AggP\_Proteasome \longrightarrow AggP\_Proteasome + PIdeath$

# $\textbf{8.1 Reaction} \; \texttt{polyQSynthesis}$

This is an irreversible reaction of one reactant forming one product.

### **Reaction equation**

$$Source \longrightarrow PolyQ \tag{6}$$

#### Reactant

Table 6: Properties of each reactant.

Id	Name	SBO
Source		

### **Product**

Table 7: Properties of each product.

Id	Name	SBO
PolyQ		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_1 = \text{ksynPolyQ} \cdot \text{Source} \cdot \text{kalive}$$
 (7)

### **8.2 Reaction** polyqProteasomeBinding

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

### **Reaction equation**

$$PolyQ + Proteasome \longrightarrow PolyQ Proteasome$$
 (8)

#### **Reactants**

Table 8: Properties of each reactant.

Id	Name	SBO
PolyQ		
Proteasome		

### **Product**

Table 9: Properties of each product.

racie 3. Freperiies of each producti			
Id	Name	SBO	
PolyQ_Proteasome			

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_2 = \text{kbinPolyQ} \cdot \text{PolyQ} \cdot \text{Proteasome} \cdot \text{kalive}$$
 (9)

# **8.3 Reaction** polyqProteasomeRelease

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

### **Reaction equation**

$$PolyQ\_Proteasome \longrightarrow PolyQ + Proteasome$$
 (10)

#### Reactant

Table 10: Properties of each reactant.

Id	Name	SBO
PolyQ_Proteasome		

### **Products**

Table 11: Properties of each product.

Id	Name	SBO
PolyQ		
Proteasome		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_3 = \text{krelPolyQ} \cdot \text{PolyQ} \cdot \text{Proteasome} \cdot \text{kalive}$$
 (11)

# **8.4 Reaction** PolyQDegradation

This is an irreversible reaction of one reactant forming one product.

### **Reaction equation**

$$PolyQ\_Proteasome \longrightarrow Proteasome \tag{12}$$

#### Reactant

Table 12: Properties of each reactant.

Id Name SBO

PolyQ\_Proteasome

#### **Product**

Table 13: Properties of each product.

Id	Name	SBO
Proteasome		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_4 = \text{kdegPolyQ} \cdot \text{PolyQ} \cdot \text{Proteasome} \cdot \text{kalive} \cdot \text{kproteff}$$
 (13)

### 8.5 Reaction mRFPuSynthesis

This is an irreversible reaction of one reactant forming one product.

### **Reaction equation**

$$Source \longrightarrow mRFPu \tag{14}$$

#### Reactant

Table 14: Properties of each reactant.

Id	Name	SBO
Source		

#### **Product**

Table 15: Properties of each product.

Id	Name	SBO
mRFPu		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_5 = \text{ksynmRFPu} \cdot \text{Source} \cdot \text{kalive}$$
 (15)

# 8.6 Reaction mRFPuProteasomeBinding

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

### **Reaction equation**

$$mRFPu + Proteasome \longrightarrow mRFPu\_Proteasome$$
 (16)

#### **Reactants**

Table 16: Properties of each reactant.

Id	Name	SBO
mRFPu		
Proteasome		

### **Product**

Table 17: Properties of each product.

Id	Name	SBO
mRFPu_Proteasome		

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_6 = \text{kbinmRFPu} \cdot \text{mRFPu} \cdot \text{Proteasome} \cdot \text{kalive}$$
 (17)

### 8.7 Reaction mRFPuProteasomeRelease

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

### **Reaction equation**

$$mRFPu\_Proteasome \longrightarrow mRFPu+Proteasome$$
 (18)

#### Reactant

Table 18: Properties of each reactant.

Id Name SBO

Iu		Ivanic	эро
mR.FP11	Proteasome		

### **Products**

Table 19: Properties of each product.

Id	Name	SBO
mRFPu		
Proteasome		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_7 = \text{krelmRFPu\_Proteasome} \cdot \text{kalive}$$
 (19)

### 8.8 Reaction mRFPuDegradation

This is an irreversible reaction of one reactant forming one product.

### **Reaction equation**

$$mRFPu\_Proteasome \longrightarrow Proteasome$$
 (20)

#### Reactant

Table 20: Properties of each reactant.

Id	Name	SBO
mRFPu_Proteasome		

#### **Product**

Table 21: Properties of each product.

Id	Name	SBO
Proteasome		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_8 = \text{kdegmRFPu} \cdot \text{mRFPu} \cdot \text{Proteasome} \cdot \text{kalive} \cdot \text{kproteff}$$
 (21)

# 8.9 Reaction PolyQAggregation1

This is an irreversible reaction of two reactants forming two products.

### **Reaction equation**

$$2 \text{PolyQ} + \text{ROS} \longrightarrow \text{AggPolyQ1} + \text{ROS}$$
 (22)

#### Reactants

Table 22: Properties of each reactant.

Id	Name	SBO
PolyQ ROS		

### **Products**

Table 23: Properties of each product.

Id	Name	SBO
AggPolyQ1 ROS		

### **Kinetic Law**

Derived unit contains undeclared units

$$\nu_9 = kaggPolyQ \cdot PolyQ \cdot (PolyQ - 1) \cdot 0.5 \cdot \frac{ROS^2}{10^2 + ROS^2} \cdot kalive \tag{23}$$

# 8.10 Reaction PolyQAggregation2

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

### **Reaction equation**

$$PolyQ + AggPolyQ1 + ROS \longrightarrow AggPolyQ2 + ROS$$
 (24)

#### **Reactants**

Table 24: Properties of each reactant.

Id	Name	SBO
PolyQ AggPolyQ1 ROS		

#### **Products**

Table 25: Properties of each product.

Id	Name	SBO
AggPolyQ2 ROS		

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{10} = kaggPolyQ \cdot PolyQ \cdot AggPolyQ1 \cdot \frac{ROS^2}{10^2 + ROS^2} \cdot kalive \tag{25}$$

# 8.11 Reaction PolyQAggregation3

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

### **Reaction equation**

$$PolyQ + AggPolyQ2 + ROS \longrightarrow AggPolyQ3 + ROS$$
 (26)

#### **Reactants**

Table 26: Properties of each reactant.

Id	Name	SBO
PolyQ AggPolyQ2 ROS		

### **Products**

Table 27: Properties of each product.

Id	Name	SBO
AggPolyQ3 ROS		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{11} = \text{kaggPolyQ} \cdot \text{PolyQ} \cdot \text{AggPolyQ2} \cdot \frac{\text{ROS}^2}{10^2 + \text{ROS}^2} \cdot \text{kalive}$$
 (27)

# 8.12 Reaction PolyQAggregation4

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

### **Reaction equation**

$$PolyQ + AggPolyQ3 + ROS \longrightarrow AggPolyQ4 + ROS$$
 (28)

#### **Reactants**

Table 28: Properties of each reactant.

Id	Name	SBO
PolyQ AggPolyQ3 ROS		

### **Products**

Table 29: Properties of each product.

Id	Name	SBO
AggPolyQ4 ROS		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{12} = \text{kaggPolyQ} \cdot \text{PolyQ} \cdot \text{AggPolyQ3} \cdot \frac{\text{ROS}^2}{10^2 + \text{ROS}^2} \cdot \text{kalive}$$
 (29)

# **8.13 Reaction** PolyQAggregation5

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

# **Reaction equation**

$$PolyQ + AggPolyQ4 + ROS \longrightarrow AggPolyQ5 + ROS$$
 (30)

#### **Reactants**

Table 30: Properties of each reactant.

Id	Name	SBO
PolyQ AggPolyQ4 ROS	:	

#### **Products**

Table 31: Properties of each product.

Id	Name	SBO
AggPolyQ5 ROS		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{13} = \text{kaggPolyQ} \cdot \text{PolyQ} \cdot \text{AggPolyQ4} \cdot \frac{\text{ROS}^2}{10^2 + \text{ROS}^2} \cdot \text{kalive}$$
 (31)

# **8.14 Reaction** PolyQDisaggregation5

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

## **Reaction equation**

$$AggPolyQ5 \longrightarrow PolyQ + AggPolyQ4 \tag{32}$$

#### Reactant

Table 32: Properties of each reactant.

Id	Name	SBO
AggPolyQ5		

#### **Products**

Table 33: Properties of each product.

Id	Name	SBO
PolyQ		
AggPolyQ4		

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{14} = \text{kdisaggPolyQ5} \cdot \text{AggPolyQ5} \cdot \text{kalive}$$
 (33)

### **8.15 Reaction** PolyQDisaggregation4

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

## **Reaction equation**

$$AggPolyQ4 \longrightarrow PolyQ + AggPolyQ3 \tag{34}$$

### Reactant

Table 34: Properties of each reactant.

Id	Name	SBO
AggPolyQ4		

### **Products**

Table 35: Properties of each product.

Id	Name	SBO
PolyQ AggPolyQ3		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{15} = \text{kdisaggPolyQ4} \cdot \text{AggPolyQ4} \cdot \text{kalive}$$
 (35)

# **8.16 Reaction** PolyQDisaggregation3

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

### **Reaction equation**

$$AggPolyQ3 \longrightarrow PolyQ + AggPolyQ2 \tag{36}$$

### Reactant

Table 36: Properties of each reactant.

Id	Name	SBO
AggPolyQ3		

### **Products**

Table 37: Properties of each product.

Id	Name	SBO
PolyQ		
AggPolyQ2		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{16} = \text{kdisaggPolyQ3} \cdot \text{AggPolyQ3} \cdot \text{kalive}$$
 (37)

# **8.17 Reaction** PolyQDisaggregation2

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

### **Reaction equation**

$$AggPolyQ2 \longrightarrow PolyQ + AggPolyQ1 \tag{38}$$

#### Reactant

Table 38: Properties of each reactant.

Id	Name	SBO
AggPolyQ2		

#### **Products**

Table 39: Properties of each product.

Id	Name	SBO
PolyQ		
AggPolyQ1		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{17} = \text{kdisaggPolyQ2} \cdot \text{AggPolyQ2} \cdot \text{kalive}$$
 (39)

# 8.18 Reaction PolyQDisaggregation1

This is an irreversible reaction of one reactant forming one product.

### **Reaction equation**

$$AggPolyQ1 \longrightarrow 2PolyQ \tag{40}$$

### Reactant

Table 40: Properties of each reactant.

Id	Name	SBO
AggPolyQ1		

### **Product**

Table 41: Properties of each product.

Id	Name	SBO
PolyQ		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{18} = \text{kdisaggPolyQ1} \cdot \text{AggPolyQ1} \cdot \text{kalive}$$
 (41)

# 8.19 Reaction PolyQInclusionFormation

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

### **Reaction equation**

$$PolyQ + AggPolyQ5 \longrightarrow 7 SeqAggP$$
 (42)

### **Reactants**

Table 42: Properties of each reactant.

Id	Name	SBO
PolyQ		
AggPolyQ5		

### **Product**

Table 43: Properties of each product.

Id	Name	SBO
SeqAggP		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{19} = \text{kaggPolyQ} \cdot \text{PolyQ} \cdot \text{AggPolyQ5} \cdot \text{kalive}$$
 (43)

### 8.20 Reaction PolyQInclusionGrowth

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

### **Reaction equation**

$$PolyQ + SeqAggP \longrightarrow 2 SeqAggP \tag{44}$$

#### **Reactants**

Table 44: Properties of each reactant.

Id	Name	SBO
PolyQ		
${\tt SeqAggP}$		

#### **Product**

Table 45: Properties of each product.

Id	Name	SBO
SeqAggP		_

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{20} = kseqPolyQ \cdot PolyQ \cdot SeqAggP \cdot kalive$$
 (45)

### 8.21 Reaction ProteasomeInhibition1

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

### **Reaction equation**

$$AggPolyQ1 + Proteasome \longrightarrow AggP\_Proteasome$$
 (46)

### **Reactants**

Table 46: Properties of each reactant.

Id	Name	SBO
AggPolyQ1 Proteasome		

#### **Product**

Table 47: Properties of each product.

Id	Name	SBO
AggP_Proteasome		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{21} = \text{kinhprot} \cdot \text{AggPolyQ1} \cdot \text{Proteasome} \cdot \text{kalive}$$
 (47)

### 8.22 Reaction ProteasomeInhibition2

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

### **Reaction equation**

$$AggPolyQ2 + Proteasome \longrightarrow AggP\_Proteasome$$
 (48)

#### **Reactants**

Table 48: Properties of each reactant.

Id	Name	SBO
AggPolyQ2 Proteasome		

#### **Product**

Table 49: Properties of each product.

ruble 15. 1 Toperties	or cach p	loudet.
Id	Name	SBO
AggP_Proteasome		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{22} = \text{kinhprot} \cdot \text{AggPolyQ2} \cdot \text{Proteasome} \cdot \text{kalive}$$
 (49)

### 8.23 Reaction ProteasomeInhibition3

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

### **Reaction equation**

$$AggPolyQ3 + Proteasome \longrightarrow AggP\_Proteasome$$
 (50)

#### Reactants

Table 50: Properties of each reactant.

Id	Name	SBO
AggPolyQ3 Proteasome		

### **Product**

Table 51: Properties of each product.

Id	Name	SBO
AggP_Proteasome		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{23} = \text{kinhprot} \cdot \text{AggPolyQ3} \cdot \text{Proteasome} \cdot \text{kalive}$$
 (51)

### 8.24 Reaction ProteasomeInhibition4

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

### **Reaction equation**

$$AggPolyQ4 + Proteasome \longrightarrow AggP\_Proteasome$$
 (52)

#### **Reactants**

Table 52: Properties of each reactant.

Name	SBO
Ī	Name

#### **Product**

Table 53: Properties of each product.

Id	Name	SBO
AggP_Proteasome		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{24} = \text{kinhprot} \cdot \text{AggPolyQ4} \cdot \text{Proteasome} \cdot \text{kalive}$$
 (53)

### 8.25 Reaction ProteasomeInhibition5

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

### **Reaction equation**

$$AggPolyQ5 + Proteasome \longrightarrow AggP\_Proteasome$$
 (54)

#### Reactants

Table 54: Properties of each reactant.

Id	Name	SBO
AggPolyQ5		

Id	Name	SBO
Proteasome		

### **Product**

Table 55: Properties of each product.

Id	Name	SBO
AggP_Proteasome		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{25} = \text{kinhprot} \cdot \text{AggPolyQ5} \cdot \text{Proteasome} \cdot \text{kalive}$$
 (55)

# **8.26 Reaction** mRFPuProteasomeSequestering

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

# **Reaction equation**

$$mRFPu\_Proteasome + SeqAggP \longrightarrow 2SeqAggP$$
 (56)

#### **Reactants**

Table 56: Properties of each reactant.

Id	Name	SBO
${\tt mRFPu\_Proteasome}$		
SeqAggP		

#### **Product**

Table 57: Properties of each product.

Id	Name	SBO
SeqAggP		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{26} = \text{kseqmRFPuProt} \cdot \text{mRFPu\_Proteasome} \cdot \text{SeqAggP} \cdot \text{kalive}$$
 (57)

# 8.27 Reaction mRFPuSequestering

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

### **Reaction equation**

$$mRFPu + SeqAggP \longrightarrow 2 SeqAggP$$
 (58)

#### **Reactants**

Table 58: Properties of each reactant.

Id	Name	SBO
mRFPu		_
${\tt SeqAggP}$		

#### **Product**

Table 59: Properties of each product.

Id	Name	SBO
SeqAggP		

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{27} = \text{kseqmRFPu} \cdot \text{mRFPu} \cdot \text{SeqAggP} \cdot \text{kalive}$$
 (59)

## 8.28 Reaction ROSgenerationBasal

This is an irreversible reaction of one reactant forming one product.

### **Reaction equation**

Source 
$$\longrightarrow$$
 ROS (60)

### Reactant

Table 60: Properties of each reactant.

Id	Name	SBO
Source		

### **Product**

Table 61: Properties of each product.

Id	Name	SBO
ROS		

### **Kinetic Law**

Derived unit contains undeclared units

$$v_{28} = \text{kgenROS} \cdot \text{Source} \cdot \text{kalive}$$
 (61)

# 8.29 Reaction ROSgenerationSmallAggPolyQ1

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

# **Reaction equation**

$$AggPolyQ1 \longrightarrow AggPolyQ1 + ROS$$
 (62)

### Reactant

Table 62: Properties of each reactant.

Id	Name	SBO
AggPolyQ1		

### **Products**

Table 63: Properties of each product.

Id	Name	SBO
AggPolyQ1		

Id	Name	SBO
ROS		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{29} = \text{kgenROSAggP} \cdot \text{AggPolyQ1} \cdot \text{kalive}$$
 (63)

### 8.30 Reaction ROSgenerationSmallAggPolyQ2

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

### **Reaction equation**

$$AggPolyQ2 \longrightarrow AggPolyQ2 + ROS$$
 (64)

#### Reactant

Table 64: Properties of each reactant.

Id	Name	SBO
AggPolyQ2		

#### **Products**

Table 65: Properties of each product.

Id	Name	SBO
AggPolyQ2 ROS		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{30} = kgenROSAggP \cdot AggPolyQ2 \cdot kalive$$
 (65)

# 8.31 Reaction ROSgenerationSmallAggPolyQ3

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

### **Reaction equation**

$$AggPolyQ3 \longrightarrow AggPolyQ3 + ROS \tag{66}$$

#### Reactant

Table 66: Properties of each reactant.

Id	Name	SBO
AggPolyQ3		

### **Products**

Table 67: Properties of each product.

Id	Name	SBO
AggPolyQ3 ROS		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{31} = kgenROSAggP \cdot AggPolyQ3 \cdot kalive$$
 (67)

# 8.32 Reaction ROSgenerationSmallAggPolyQ4

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

### **Reaction equation**

$$AggPolyQ4 \longrightarrow AggPolyQ4 + ROS \tag{68}$$

#### Reactant

Table 68: Properties of each reactant.

Id	Name	SBO
AggPolyQ4		

#### **Products**

Table 69: Properties of each product.

Id	Name	SBO
AggPolyQ4 ROS		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{32} = kgenROSAggP \cdot AggPolyQ4 \cdot kalive$$
 (69)

# 8.33 Reaction ROSgenerationSmallAggPolyQ5

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

### **Reaction equation**

$$AggPolyQ5 \longrightarrow AggPolyQ5 + ROS$$
 (70)

### Reactant

Table 70: Properties of each reactant.

Id	Name	SBO
AggPolyQ5		

### **Products**

Table 71: Properties of each product.

Id	Name	SBO
AggPolyQ5 ROS		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{33} = \text{kgenROSAggP} \cdot \text{AggPolyQ5} \cdot \text{kalive}$$
 (71)

# **8.34 Reaction** ROSgenerationAggPProteasome

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

### **Reaction equation**

$$AggP\_Proteasome \longrightarrow AggP\_Proteasome + ROS$$
 (72)

#### Reactant

Table 72: Properties of each reactant.

Id	Name	SBO
AggP_Proteasome		

### **Products**

Table 73: Properties of each product.

Id	Name	SBO
AggP_Proteasome ROS		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{34} = \text{kgenROSAggP} \cdot \text{AggP\_Proteasome} \cdot \text{kalive}$$
 (73)

### 8.35 Reaction ROSremoval

This is an irreversible reaction of one reactant forming one product.

### **Reaction equation**

$$ROS \longrightarrow Sink$$
 (74)

#### Reactant

Table 74: Properties of each reactant.

Id	Name	SBO
ROS		

Table 75: Properties of each product.

Id	Name	SBO
Sink		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{35} = \text{kremROS} \cdot \text{ROS} \cdot \text{kalive}$$
 (75)

# 8.36 Reaction p38activation

This is an irreversible reaction of two reactants forming two products.

# **Reaction equation**

$$ROS + p38 \longrightarrow ROS + p38\_P \tag{76}$$

### **Reactants**

Table 76: Properties of each reactant.

Id	Name	SBO
ROS		
p38		

### **Products**

Table 77: Properties of each product.

Id	Name	SBO
ROS		
p38_P		

# **Kinetic Law**

$$v_{36} = \text{kactp38} \cdot \text{ROS} \cdot \text{p38} \cdot \text{kalive}$$
 (77)

# **8.37 Reaction** p38inactivation

This is an irreversible reaction of one reactant forming one product.

# **Reaction equation**

$$p38\_P \longrightarrow p38 \tag{78}$$

### Reactant

Table 78: Properties of each reactant.

Id	Name	SBO
p38_P		

# **Product**

Table 79: Properties of each product.

Id	Name	SBO
p38		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{37} = \text{kinactp38} \cdot \text{p38} \cdot \text{P} \cdot \text{kalive}$$
 (79)

# **8.38 Reaction** AggP\_ProteasomeSequestering

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

# **Reaction equation**

$$AggP\_Proteasome + SeqAggP \longrightarrow 2 SeqAggP$$
 (80)

### **Reactants**

Table 80: Properties of each reactant.

Id	Name	SBO
AggP_Proteasome SeqAggP		

Table 81: Properties of each product.

Id	Name	SBO
SeqAggP		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{38} = \text{kseqAggPProt} \cdot \text{AggP\_Proteasome} \cdot \text{SeqAggP} \cdot \text{kalive}$$
 (81)

# 8.39 Reaction PolyQ\_ProteasomeSequestering

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

# **Reaction equation**

$$PolyQ\_Proteasome + SeqAggP \longrightarrow 2SeqAggP$$
 (82)

### **Reactants**

Table 82: Properties of each reactant.

Id	Name	SBO
PolyQ_Proteasome SeqAggP		

### **Product**

Table 83: Properties of each product.

Id	Name	SBO
SeqAggP		

### **Kinetic Law**

$$v_{39} = \text{kseqPolyQProt} \cdot \text{PolyQ\_Proteasome} \cdot \text{SeqAggP} \cdot \text{kalive}$$
 (83)

# **8.40 Reaction** MisP\_ProteasomeSequestering

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

# **Reaction equation**

$$MisP\_Proteasome + SeqAggP \longrightarrow 2 SeqAggP$$
 (84)

### **Reactants**

Table 84: Properties of each reactant.

Id	Name	SBO
MisP_Proteasome		
${\tt SeqAggP}$		

#### **Product**

Table 85: Properties of each product.

Id	Name	SBO
SeqAggP		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{40} = \text{kseqMisPProt} \cdot \text{MisP\_Proteasome} \cdot \text{SeqAggP} \cdot \text{kalive}$$
 (85)

# 8.41 Reaction ProteinSynthesis

This is an irreversible reaction of one reactant forming one product.

# **Reaction equation**

Source 
$$\longrightarrow$$
 NatP (86)

#### Reactant

Table 86: Properties of each reactant.

Id	Name	SBO
Source		

Table 87: Properties of each product.

Id	Name	SBO
NatP		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{41} = \text{ksynNatP} \cdot \text{Source} \cdot \text{kalive}$$
 (87)

# 8.42 Reaction Misfolding

This is an irreversible reaction of two reactants forming two products.

# **Reaction equation**

$$NatP + ROS \longrightarrow MisP + ROS$$
 (88)

### **Reactants**

Table 88: Properties of each reactant.

Id	Name	SBO
NatP		·
ROS		

### **Products**

Table 89: Properties of each product.

Id	Name	SBO
MisP		
ROS		

# **Kinetic Law**

$$v_{42} = \text{kmisfold} \cdot \text{NatP} \cdot \text{ROS} \cdot \text{kalive}$$
 (89)

# 8.43 Reaction Refolding

This is an irreversible reaction of one reactant forming one product.

# **Reaction equation**

$$MisP \longrightarrow NatP$$
 (90)

### Reactant

Table 90: Properties of each reactant.

Id	Name	SBO
MisP		

# **Product**

Table 91: Properties of each product.

Id	Name	SBO
NatP		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{43} = \text{krefold} \cdot \text{MisP} \cdot \text{kalive}$$
 (91)

# **8.44 Reaction** MisP\_ProteasomeBinding

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

# **Reaction equation**

$$MisP + Proteasome \longrightarrow MisP\_Proteasome$$
 (92)

### **Reactants**

Table 92: Properties of each reactant.

Id	Name	SBO
MisP		_
Proteasome		

Table 93: Properties of each product.

ruere yet rreperiors	or caren p	
Id	Name	SBO
MisP_Proteasome		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{44} = \text{kbinMisPProt} \cdot \text{MisP} \cdot \text{Proteasome} \cdot \text{kalive}$$
 (93)

# 8.45 Reaction MisP\_ProteasomeRelease

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

# **Reaction equation**

$$MisP\_Proteasome \longrightarrow MisP+Proteasome$$
 (94)

### Reactant

Table 94: Properties of each reactant.

Id	Name	SBO
MisP_Proteasome		

# **Products**

Table 95: Properties of each product.

Id	Name	SBO
MisP		
Proteasome		

### **Kinetic Law**

$$v_{45} = \text{krelMisPProt} \cdot \text{MisP\_Proteasome} \cdot \text{kalive}$$
 (95)

# **8.46 Reaction** MisP\_Degradation

This is an irreversible reaction of one reactant forming one product.

# **Reaction equation**

$$MisP\_Proteasome \longrightarrow Proteasome$$
 (96)

### Reactant

Table 96: Properties of each reactant.

Id Name SBO

MisP\_Proteasome

# **Product**

Table 97: Properties of each product.

Id	Name	SBO
Proteasome		

# **Kinetic Law**

Derived unit contains undeclared units

$$v_{46} = \text{kdegMisP} \cdot \text{MisP\_Proteasome} \cdot \text{kalive} \cdot \text{kproteff}$$
 (97)

# 8.47 Reaction MisP\_Aggregation1

This is an irreversible reaction of one reactant forming one product.

# **Reaction equation**

$$2MisP \longrightarrow AggP1$$
 (98)

### Reactant

Table 98: Properties of each reactant.

Id	Name	SBO
MisP		

Table 99: Properties of each product.

Id	Name	SBO
AggP1		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{47} = \text{kaggMisP} \cdot \text{MisP} \cdot (\text{MisP} - 1) \cdot 0.5 \cdot \text{kalive}$$
 (99)

# **8.48 Reaction** MisP\_Aggregation2

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

# **Reaction equation**

$$MisP + AggP1 \longrightarrow AggP2 \tag{100}$$

# **Reactants**

Table 100: Properties of each reactant.

Id	Name	SBO
MisP		
AggP1		

# **Product**

Table 101: Properties of each product.

Id	Name	SBO
AggP2		

# **Kinetic Law**

$$v_{48} = \text{kagg2MisP} \cdot \text{MisP} \cdot \text{AggP1} \cdot \text{kalive}$$
 (101)

# **8.49 Reaction** MisP\_Aggregation3

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

# **Reaction equation**

$$MisP + AggP2 \longrightarrow AggP3 \tag{102}$$

### **Reactants**

Table 102: Properties of each reactant.

Id	Name	SBO
MisP		
AggP2		

#### **Product**

Table 103: Properties of each product.

Id	Name	SBO
AggP3		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{49} = \text{kagg2MisP} \cdot \text{MisP} \cdot \text{AggP2} \cdot \text{kalive}$$
 (103)

# 8.50 Reaction MisP\_Aggregation4

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

# **Reaction equation**

$$MisP + AggP3 \longrightarrow AggP4 \tag{104}$$

#### **Reactants**

Table 104: Properties of each reactant.

Id	Name	SBO
MisP		

Id	Name	SBO
AggP3		

Table 105: Properties of each product.

Id	Name	SBO
AggP4		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{50} = \text{kagg2MisP} \cdot \text{MisP} \cdot \text{AggP3} \cdot \text{kalive}$$
 (105)

# **8.51 Reaction MisP\_Aggregation5**

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

# **Reaction equation**

$$MisP + AggP4 \longrightarrow AggP5 \tag{106}$$

### **Reactants**

Table 106: Properties of each reactant.

Id	Name	SBO
MisP		
AggP4		

### **Product**

Table 107: Properties of each product.

Id	Name	SBO
AggP5		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{51} = \text{kagg2MisP} \cdot \text{MisP} \cdot \text{AggP4} \cdot \text{kalive}$$
 (107)

# **8.52 Reaction** MisP\_Disaggregation1

This is an irreversible reaction of one reactant forming one product.

# **Reaction equation**

$$AggP1 \longrightarrow 2MisP \tag{108}$$

# Reactant

Table 108: Properties of each reactant.

Id	Name	SBO
AggP1		

### **Product**

Table 109: Properties of each product.

Id	Name	SBO
MisP		

# **Kinetic Law**

Derived unit contains undeclared units

$$v_{52} = \text{kdisaggMisP1} \cdot \text{AggP1} \cdot \text{kalive}$$
 (109)

# **8.53 Reaction** MisP\_Disaggregation2

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

# **Reaction equation**

$$AggP2 \longrightarrow MisP + AggP1 \tag{110}$$

# Reactant

Table 110: Properties of each reactant.

Id	Name	SBO
AggP2		

Table 111: Properties of each product.

Id	Name	SBO
MisP		
AggP1		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{53} = \text{kdisaggMisP2} \cdot \text{AggP2} \cdot \text{kalive}$$
 (111)

# **8.54 Reaction** MisP\_Disaggregation3

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

# **Reaction equation**

$$AggP3 \longrightarrow MisP + AggP2 \tag{112}$$

### Reactant

Table 112: Properties of each reactant.

Id	Name	SBO
AggP3		

# **Products**

Table 113: Properties of each product.

Id	Name	SBO
MisP		
AggP2		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{54} = \text{kdisaggMisP3} \cdot \text{AggP3} \cdot \text{kalive}$$
 (113)

# 8.55 Reaction MisP\_Disaggregation4

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

# **Reaction equation**

$$AggP4 \longrightarrow MisP + AggP3 \tag{114}$$

### Reactant

Table 114: Properties of each reactant.

Id	Name	SBO
AggP4		

### **Products**

Table 115: Properties of each product.

Id	Name	SBO
MisP		
AggP3		

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{55} = \text{kdisaggMisP4} \cdot \text{AggP4} \cdot \text{kalive}$$
 (115)

# **8.56 Reaction** MisP\_Disaggregation5

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

# **Reaction equation**

$$AggP5 \longrightarrow MisP + AggP4 \tag{116}$$

# Reactant

Table 116: Properties of each reactant.

Id	Name	SBO
AggP5		

# **Products**

Table 117: Properties of each product.

Id	Name	SBO
MisP		
AggP4		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{56} = \text{kdisaggMisP5} \cdot \text{AggP5} \cdot \text{kalive}$$
 (117)

# **8.57 Reaction** MisP\_InclusionFormation

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

# **Reaction equation**

$$MisP + AggP5 \longrightarrow 7 SeqAggP$$
 (118)

### **Reactants**

Table 118: Properties of each reactant.

Id	Name	SBO
MisP		
AggP5		

### **Product**

Table 119: Properties of each product.

Id	Name	SBO
SeqAggP		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{57} = \text{kagg2MisP} \cdot \text{MisP} \cdot \text{AggP5} \cdot \text{kalive}$$
 (119)

# 8.58 Reaction MisPInclusionGrowth

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

# **Reaction equation**

$$MisP + SeqAggP \longrightarrow 2 SeqAggP \tag{120}$$

### **Reactants**

Table 120: Properties of each reactant.

Id	Name	SBO
MisP		
${\tt SeqAggP}$		

### **Product**

Table 121: Properties of each product.

Id	Name	SBO
SeqAggP		

### **Kinetic Law**

Derived unit contains undeclared units

$$v_{58} = \text{kseqMisP} \cdot \text{MisP} \cdot \text{SeqAggP} \cdot \text{kalive}$$
 (121)

# **8.59 Reaction** ProteasomeInhibitionAggP1

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

# **Reaction equation**

$$AggP1 + Proteasome \longrightarrow AggP\_Proteasome$$
 (122)

### **Reactants**

Table 122: Properties of each reactant.

Id	Name	SBO
AggP1 Proteasome		

### **Product**

Table 123: Properties of each product.

Id	Name	SBO
AggP_Proteasome		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{59} = \text{kinhprot} \cdot \text{AggP1} \cdot \text{Proteasome} \cdot \text{kalive}$$
 (123)

# **8.60 Reaction** ProteasomeInhibitionAggP2

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

# **Reaction equation**

$$AggP2 + Proteasome \longrightarrow AggP\_Proteasome$$
 (124)

#### **Reactants**

Table 124: Properties of each reactant.

Id	Name	SBO
AggP2 Proteasome		

Table 125: Properties of each product.

Tueste 128: 11eperties	or cach p	or o a a c c .
Id	Name	SBO
AggP_Proteasome		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{60} = \text{kinhprot} \cdot \text{AggP2} \cdot \text{Proteasome} \cdot \text{kalive}$$
 (125)

# 8.61 Reaction ProteasomeInhibitionAggP3

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

# **Reaction equation**

$$AggP3 + Proteasome \longrightarrow AggP\_Proteasome$$
 (126)

### **Reactants**

Table 126: Properties of each reactant.

Id	Name	SBO
AggP3 Proteasome		

### **Product**

Table 127: Properties of each product.

Id	Name	SBO
AggP_Proteasome		

### **Kinetic Law**

$$v_{61} = \text{kinhprot} \cdot \text{AggP3} \cdot \text{Proteasome} \cdot \text{kalive}$$
 (127)

# 8.62 Reaction ProteasomeInhibitionAggP4

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

# **Reaction equation**

$$AggP4 + Proteasome \longrightarrow AggP\_Proteasome$$
 (128)

### **Reactants**

Table 128: Properties of each reactant.

Id	Name	SBO
AggP4 Proteasome		

#### **Product**

Table 129: Properties of each product.

Id	Name	SBO
AggP_Proteasome		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{62} = \text{kinhprot} \cdot \text{AggP4} \cdot \text{Proteasome} \cdot \text{kalive}$$
 (129)

# 8.63 Reaction ProteasomeInhibitionAggP5

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

# **Reaction equation**

$$AggP5 + Proteasome \longrightarrow AggP\_Proteasome$$
 (130)

#### **Reactants**

Table 130: Properties of each reactant.

Id	Name	
AggP5		

Id	Name	SBO
Proteasome		

Table 131: Properties of each product.

Tuble 131: 1 Toperties of each product.		
Id	Name	SBO
AggP_Proteasome		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{63} = \text{kinhprot} \cdot \text{AggP5} \cdot \text{Proteasome} \cdot \text{kalive}$$
 (131)

# **8.64 Reaction** ROSgenerationSmallAggP1

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

# **Reaction equation**

$$AggP1 \longrightarrow AggP1 + ROS \tag{132}$$

### Reactant

Table 132: Properties of each reactant.

Id	Name	SBO
AggP1		

# **Products**

Table 133: Properties of each product.

Id	Name	SBO
AggP1 ROS		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{64} = \text{kgenROSAggP} \cdot \text{AggP1} \cdot \text{kalive}$$
 (133)

# 8.65 Reaction ROSgenerationSmallAggP2

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

# **Reaction equation**

$$AggP2 \longrightarrow AggP2 + ROS \tag{134}$$

### Reactant

Table 134: Properties of each reactant.

Id	Name	SBO
AggP2		

### **Products**

Table 135: Properties of each product.

Id	Name	SBO
AggP2 ROS		

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{65} = kgenROSAggP \cdot AggP2 \cdot kalive$$
 (135)

# 8.66 Reaction ROSgenerationSmallAggP3

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

# **Reaction equation**

$$AggP3 \longrightarrow AggP3 + ROS \tag{136}$$

# Reactant

Table 136: Properties of each reactant.

Id	Name	SBO
AggP3		

# **Products**

Table 137: Properties of each product.

Id	Name	SBO
AggP3 ROS		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{66} = \text{kgenROSAggP} \cdot \text{AggP3} \cdot \text{kalive}$$
 (137)

# 8.67 Reaction ROSgenerationSmallAggP4

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

# **Reaction equation**

$$AggP4 \longrightarrow AggP4 + ROS \tag{138}$$

### Reactant

Table 138: Properties of each reactant.

Id	Name	SBO
AggP4		

### **Products**

Table 139: Properties of each product.

Id	Name	SBO
AggP4 ROS		

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{67} = \text{kgenROSAggP} \cdot \text{AggP4} \cdot \text{kalive}$$
 (139)

# 8.68 Reaction ROSgenerationSmallAggP5

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

# **Reaction equation**

$$AggP5 \longrightarrow AggP5 + ROS \tag{140}$$

# Reactant

Table 140: Properties of each reactant.

Id	Name	SBO
AggP5		

### **Products**

Table 141: Properties of each product.

Id	Name	SBO
AggP5 ROS		

# **Kinetic Law**

$$v_{68} = \text{kgenROSAggP} \cdot \text{AggP5} \cdot \text{kalive}$$
 (141)

# 8.69 Reaction p38\_P\_ROS\_Generation

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

# **Reaction equation**

$$p38\_P \longrightarrow p38\_P + ROS$$
 (142)

### Reactant

Table 142: Properties of each reactant.

Id	Name	SBO
p38_P		

# **Products**

Table 143: Properties of each product.

Id	Name	SBO
p38_P ROS		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{69} = \text{kgenROSp38} \cdot \text{p38} \cdot \text{P} \cdot \text{kp38act} \cdot \text{kalive}$$
 (143)

# 8.70 Reaction ROSgenerationSeqAggP

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

# **Reaction equation**

$$SeqAggP \longrightarrow SeqAggP + ROS \tag{144}$$

#### Reactant

Table 144: Properties of each reactant.

Id	Name	SBO
SeqAggP		

Table 145: Properties of each product.

Id	Name	SBO
SeqAggP ROS		

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{70} = \text{kgenROSSeqAggP} \cdot \text{SeqAggP} \cdot \text{kalive}$$
 (145)

# **8.71 Reaction** P38DeathPathway

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

# **Reaction equation**

$$p38\_P \longrightarrow p38\_P + p38death$$
 (146)

#### Reactant

Table 146: Properties of each reactant.

Id	Name	SBO
p38_P		

### **Products**

Table 147: Properties of each product.

Id	Name	SBO
p38_P		
p38death		

# **Kinetic Law**

$$v_{71} = \text{kp38death} \cdot \text{p38} \cdot \text{P} \cdot \text{kalive} \cdot \text{kp38act}$$
 (147)

# 8.72 Reaction PIDeathPathway

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

### **Reaction equation**

$$AggP\_Proteasome \longrightarrow AggP\_Proteasome + PIdeath$$
 (148)

#### Reactant

Table 148: Properties of each reactant.

Id Name SBO

AggP\_Proteasome

#### **Products**

Table 149: Properties of each product.

Id Name SBO

AggP\_Proteasome
PIdeath

### **Kinetic Law**

Derived unit contains undeclared units

$$v_{72} = \text{kPIdeath} \cdot \text{AggP\_Proteasome} \cdot \text{kalive}$$
 (149)

# 9 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.

### 9.1 Species PolyQ

#### Initial amount 1000 item

This species takes part in 15 reactions (as a reactant in polyqProteasomeBinding, PolyQAggregation1, PolyQAggregation2, PolyQAggregation3, PolyQAggregation4, PolyQAggregation5, PolyQInclusionFor PolyQInclusionGrowth and as a product in polyQSynthesis, polyqProteasomeRelease, PolyQDisaggregation5, PolyQDisaggregation4, PolyQDisaggregation3, PolyQDisaggregation2, PolyQDisaggregation1).

$$\frac{d}{dt} \text{PolyQ} = v_1 + v_3 + v_{14} + v_{15} + v_{16} + v_{17} + 2v_{18} - v_2 
-2v_9 - v_{10} - v_{11} - v_{12} - v_{13} - v_{19} - v_{20}$$
(150)

# 9.2 Species Proteasome

#### Initial amount 1000 item

This species takes part in 19 reactions (as a reactant in polyqProteasomeBinding, mRFPuProteasomeBinding, ProteasomeInhibition1, ProteasomeInhibition2, ProteasomeInhibition3, ProteasomeInhibition4, ProteasomeInhibition5, Misp\_ProteasomeBinding, ProteasomeInhibitionAggP1, ProteasomeInhibiti ProteasomeInhibitionAggP3, ProteasomeInhibitionAggP4, ProteasomeInhibitionAggP5 and as a product in polyqProteasomeRelease, PolyQDegradation, mRFPuProteasomeRelease, mRFPuDegradation, Misp\_ProteasomeRelease, Misp\_Degradation).

$$\frac{d}{dt} \text{Proteasome} = v_3 + v_4 + v_7 + v_8 + v_{45} + v_{46} - v_2 - v_6 - v_{21} - v_{22} - v_{60} - v_{21} - v_{22} - v_{60} - v_{61} - v_{62} - v_{63}$$

$$(151)$$

### 9.3 Species NatP

#### Initial amount 19500 item

This species takes part in three reactions (as a reactant in Misfolding and as a product in ProteinSynthesis, Refolding).

$$\frac{d}{dt}NatP = |v_{41}| + |v_{43}| - |v_{42}|$$
 (152)

### 9.4 Species MisP

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

This species takes part in 16 reactions (as a reactant in Refolding, MisP\_ProteasomeBinding, MisP\_Aggregation1, MisP\_Aggregation2, MisP\_Aggregation3, MisP\_Aggregation4, MisP\_Aggregation5, MisP\_InclusionFormation, MisPInclusionGrowth and as a product in

Misfolding, MisP\_ProteasomeRelease, MisP\_Disaggregation1, MisP\_Disaggregation2, MisP\_Disaggregation3, MisP\_Disaggregation4, MisP\_Disaggregation5).

$$\frac{d}{dt}MisP = v_{42} + v_{45} + 2 v_{52} + v_{53} + v_{54} + v_{55} + v_{56} - v_{43} - v_{44} - 2 v_{47} - v_{48} - v_{49} - v_{50} - v_{51} - v_{57} - v_{58}$$
(153)

### 9.5 Species MisP\_Proteasome

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

This species takes part in four reactions (as a reactant in MisP\_ProteasomeSequestering, MisP\_ProteasomeRelease, MisP\_Degradation and as a product in MisP\_ProteasomeBinding).

$$\frac{d}{dt} \text{MisP\_Proteasome} = |v_{44}| - |v_{40}| - |v_{45}| - |v_{46}|$$
 (154)

# 9.6 Species AggP1

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

This species takes part in seven reactions (as a reactant in MisP\_Aggregation2, MisP\_Disaggregation1, ProteasomeInhibitionAggP1, ROSgenerationSmallAggP1 and as a product in MisP\_Aggregation1, MisP\_Disaggregation2, ROSgenerationSmallAggP1).

$$\frac{d}{dt}AggP1 = |v_{47}| + |v_{53}| + |v_{64}| - |v_{48}| - |v_{52}| - |v_{59}| - |v_{64}|$$
(155)

# 9.7 Species AggP2

### **Initial amount** 0 item

This species takes part in seven reactions (as a reactant in MisP\_Aggregation3, MisP\_Disaggregation2, ProteasomeInhibitionAggP2, ROSgenerationSmallAggP2 and as a product in MisP\_Aggregation2, MisP\_Disaggregation3, ROSgenerationSmallAggP2).

$$\frac{d}{dt}AggP2 = v_{48} + v_{54} + v_{65} - v_{49} - v_{53} - v_{60} - v_{65}$$
(156)

### 9.8 Species AggP3

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

This species takes part in seven reactions (as a reactant in MisP\_Aggregation4, MisP\_Disaggregation3, ProteasomeInhibitionAggP3, ROSgenerationSmallAggP3 and as a product in MisP\_Aggregation3, MisP\_Disaggregation4, ROSgenerationSmallAggP3).

$$\frac{d}{dt}AggP3 = v_{49} + v_{55} + v_{66} - v_{50} - v_{54} - v_{61} - v_{66}$$
(157)

### 9.9 Species AggP4

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

This species takes part in seven reactions (as a reactant in MisP\_Aggregation5, MisP\_Disaggregation4, ProteasomeInhibitionAggP4, ROSgenerationSmallAggP4 and as a product in MisP\_Aggregation4, MisP\_Disaggregation5, ROSgenerationSmallAggP4).

$$\frac{\mathrm{d}}{\mathrm{d}t} AggP4 = |v_{50}| + |v_{56}| + |v_{67}| - |v_{51}| - |v_{55}| - |v_{62}| - |v_{67}|$$
(158)

### 9.10 Species AggP5

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

This species takes part in six reactions (as a reactant in MisP\_Disaggregation5, MisP\_InclusionFormation, ProteasomeInhibitionAggP5, ROSgenerationSmallAggP5 and as a product in MisP\_Aggregation5, ROSgenerationSmallAggP5).

$$\frac{d}{dt}AggP5 = |v_{51}| + |v_{68}| - |v_{56}| - |v_{57}| - |v_{63}| - |v_{68}|$$
(159)

# 9.11 Species AggPolyQ1

# **Initial amount** 0 item

This species takes part in seven reactions (as a reactant in PolyQAggregation2, PolyQDisaggregation1, ProteasomeInhibition1, ROSgenerationSmallAggPolyQ1 and as a product in PolyQAggregation1, PolyQDisaggregation2, ROSgenerationSmallAggPolyQ1).

$$\frac{d}{dt}AggPolyQ1 = v_9 + v_{17} + v_{29} - v_{10} - v_{18} - v_{21} - v_{29}$$
(160)

# 9.12 Species AggPolyQ2

# **Initial amount** 0 item

This species takes part in seven reactions (as a reactant in PolyQAggregation3, PolyQDisaggregation2, ProteasomeInhibition2, ROSgenerationSmallAggPolyQ2 and as a product in PolyQAggregation2, PolyQDisaggregation3, ROSgenerationSmallAggPolyQ2).

$$\frac{d}{dt}AggPolyQ2 = |v_{10}| + |v_{16}| + |v_{30}| - |v_{11}| - |v_{17}| - |v_{22}| - |v_{30}|$$
(161)

### 9.13 Species AggPolyQ3

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

This species takes part in seven reactions (as a reactant in PolyQAggregation4, PolyQDisaggregation3, ProteasomeInhibition3, ROSgenerationSmallAggPolyQ3 and as a product in PolyQAggregation3, PolyQDisaggregation4, ROSgenerationSmallAggPolyQ3).

$$\frac{d}{dt}AggPolyQ3 = |v_{11}| + |v_{15}| + |v_{31}| - |v_{12}| - |v_{16}| - |v_{23}| - |v_{31}|$$
(162)

### 9.14 Species AggPolyQ4

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

This species takes part in seven reactions (as a reactant in PolyQAggregation5, PolyQDisaggregation4, ProteasomeInhibition4, ROSgenerationSmallAggPolyQ4 and as a product in PolyQAggregation4, PolyQDisaggregation5, ROSgenerationSmallAggPolyQ4).

$$\frac{d}{dt} AggPolyQ4 = v_{12} + v_{14} + v_{32} - v_{13} - v_{15} - v_{24} - v_{32}$$
(163)

# 9.15 Species AggPolyQ5

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

This species takes part in six reactions (as a reactant in PolyQDisaggregation5, PolyQInclusionFormation, ProteasomeInhibition5, ROSgenerationSmallAggPolyQ5 and as a product in PolyQAggregation5, ROSgenerationSmallAggPolyQ5).

$$\frac{d}{dt}AggPolyQ5 = |v_{13}| + |v_{33}| - |v_{14}| - |v_{19}| - |v_{25}| - |v_{33}|$$
(164)

# 9.16 Species SeqAggP

### **Initial amount** 0 item

This species takes part in 18 reactions (as a reactant in PolyQInclusionGrowth, mRFPuProteasomeSequestering mRFPuSequestering, AggP\_ProteasomeSequestering, PolyQ\_ProteasomeSequestering, MisP\_ProteasomeSequestering, MisPInclusionGrowth, ROSgenerationSeqAggP and as a product in PolyQInclusionFormation, PolyQInclusionGrowth, mRFPuProteasomeSequestering, mRFPuSequestering, AggP\_ProteasomeSequestering, PolyQ\_ProteasomeSequestering, MisP\_ProteasomeSequestering, MisP\_InclusionFormation, MisPInclusionGrowth, ROSgenerationSeqAggP\_ProteasomeSequestering, MisP\_InclusionGrowth, ROS

$$\frac{d}{dt} \text{SeqAggP} = 7 v_{19} + 2 v_{20} + 2 v_{26} + 2 v_{27} + 2 v_{38} + 2 v_{39} + 2 v_{40} + 7 v_{57} + 2 v_{58} + v_{70} - v_{20} - v_{26} - v_{27} - v_{38} - v_{39} - v_{40} - v_{58} - v_{70}$$
(165)

### 9.17 Species AggP\_Proteasome

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

This species takes part in 15 reactions (as a reactant in ROSgenerationAggPProteasome, AggP\_ProteasomeSequestering, PIDeathPathway and as a product in ProteasomeInhibition1, ProteasomeInhibition2, ProteasomeInhibition3, ProteasomeInhibition4, ProteasomeInhibition5, ROSgenerationAggPProteasome, ProteasomeInhibitionAggP1, ProteasomeInhibitionAggP2, ProteasomeInhibitionAggP3, ProteasomeInhibitionAggP4, ProteasomeInhibitionAggP5, PIDeathPathway).

$$\frac{d}{dt} AggP\_Proteasome = v_{21} + v_{22} + v_{23} + v_{24} + v_{25} + v_{34} + v_{59} + v_{60}$$

$$+ v_{61} + v_{62} + v_{63} + v_{72} - v_{34} - v_{38} - v_{72}$$
(166)

# 9.18 Species mRFPu

#### Initial amount 300 item

This species takes part in four reactions (as a reactant in mRFPuProteasomeBinding, mRFPuSequestering and as a product in mRFPuSynthesis, mRFPuProteasomeRelease).

$$\frac{d}{dt} mRFPu = |v_5| + |v_7| - |v_6| - |v_{27}|$$
(167)

# 9.19 Species mRFPu\_Proteasome

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

This species takes part in four reactions (as a reactant in mRFPuProteasomeRelease, mRFPuDegradation, mRFPuProteasomeSequestering and as a product in mRFPuProteasomeBinding).

$$\frac{d}{dt} mRFPu\_Proteasome = v_6 - v_7 - v_8 - v_{26}$$
 (168)

# 9.20 Species PolyQ\_Proteasome

### **Initial amount** 0 item

This species takes part in four reactions (as a reactant in polyqProteasomeRelease, PolyQDegradation, PolyQ\_ProteasomeSequestering and as a product in polyqProteasomeBinding).

$$\frac{\mathrm{d}}{\mathrm{d}t} \text{PolyQ\_Proteasome} = |v_2| - |v_3| - |v_4| - |v_{39}| \tag{169}$$

### 9.21 Species ROS

#### **Initial amount** 10 item

This species takes part in 29 reactions (as a reactant in PolyQAggregation1, PolyQAggregation2, PolyQAggregation3, PolyQAggregation4, PolyQAggregation5, ROSremoval, p38activation, Misfolding and as a product in PolyQAggregation1, PolyQAggregation2, PolyQAggregation3, PolyQAggregation4, PolyQAggregation5, ROSgenerationBasal, ROSgenerationSmallAggPolyQ1, ROSgenerationSmallAggPolyQ2, ROSgenerationSmallAggPolyQ3, ROSgenerationSmallAggPolyQ4, ROSgenerationSmallAggPolyQ5, ROSgenerationAggPProteasome, p38activation, Misfolding, ROSgenerationSmallAggP1, ROSgenerationSmallAggP2, ROSgenerationSmallAggP3, ROSgenerationSmallAggP5, p38\_P\_ROS\_Generation, ROSgenerationSeqAggP).

$$\frac{d}{dt}ROS = v_9 + v_{10} + v_{11} + v_{12} + v_{13} + v_{28} + v_{29} + v_{30} + v_{31} + v_{32} + v_{33} + v_{34} + v_{36} + v_{42} + v_{64} + v_{65} + v_{66} + v_{67} + v_{68} + v_{69} + v_{70} - v_9 - v_{10} - v_{11} - v_{12} - v_{13} - v_{35} - v_{36} - v_{42}$$

$$(170)$$

# **9.22 Species** p38\_P

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

This species takes part in six reactions (as a reactant in p38inactivation, p38\_P\_ROS\_Generation, P38DeathPathway and as a product in p38activation, p38\_P\_ROS\_Generation, P38DeathPathway).

$$\frac{d}{dt}p38.P = |v_{36}| + |v_{69}| + |v_{71}| - |v_{37}| - |v_{69}| - |v_{71}|$$
(171)

### **9.23 Species** p38

### Initial amount 100 item

This species takes part in two reactions (as a reactant in p38activation and as a product in p38inactivation).

$$\frac{d}{dt}p38 = |v_{37}| - |v_{36}| \tag{172}$$

#### **9.24 Species** Source

#### **Initial amount** 1 item

This species takes part in four reactions (as a reactant in polyQSynthesis, mRFPuSynthesis, ROSgenerationBasal, ProteinSynthesis), which do not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Source} = 0\tag{173}$$

# 9.25 Species Sink

#### **Initial amount** 1 item

This species takes part in one reaction (as a product in ROSremoval), which does not influence its rate of change because this constant species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Sink} = 0\tag{174}$$

# 9.26 Species p38death

### **Initial amount** 0 item

This species takes part in one reaction (as a product in P38DeathPathway).

$$\frac{\mathrm{d}}{\mathrm{d}t} p38 \mathrm{death} = v_{71} \tag{175}$$

# 9.27 Species PIdeath

### **Initial amount** 0 item

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

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{PIdeath} = v_{72} \tag{176}$$

 $\mathfrak{BML2}^{d}$  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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