# **SBML Model Report**

# Model name: "Sasagawa2005\_MAPK"



May 5, 2016

#### 1 General Overview

This is a document in SBML Level 2 Version 1 format. This model was created by the following two authors: Lu Li<sup>1</sup> and Shinya Kuroda<sup>2</sup> at December 21<sup>st</sup> 2005 at 10:59 a.m. and last time modified at April fourth 2014 at 1:27 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         | 2        |
| species types     | 0        | species              | 99       |
| events            | 0        | constraints          | 0        |
| reactions         | 150      | function definitions | 0        |
| global parameters | 0        | unit definitions     | 1        |
| rules             | 0        | initial assignments  | 0        |

#### **Model Notes**

This a model from the article:

Prediction and validation of the distinct dynamics of transient and sustained ERK activation.

Sasagawa S, Ozaki Y, Fujita K, Kuroda S <u>Nat. Cell Biol.</u>[2005 Apr; Volume: 7 (Issue: 4 )]: 365-73 15793571,

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#### **Abstract:**

To elucidate the hidden dynamics of extracellular-signal-regulated kinase (ERK) signalling networks, we developed a simulation model of ERK signalling networks by constraining in silico dynamics based on in vivo dynamics in PC12 cells. We predicted and validated that transient ERK activation depends on rapid increases of epidermal growth factor and nerve growth factor (NGF) but not on their final concentrations, whereas sustained ERK activation depends on the final concentration of NGF but not on the temporal rate of increase. These ERK dynamics depend on Ras and Rap1 dynamics, the inactivation processes of which are growth-factor-dependent and -independent, respectively. Therefore, the Ras and Rap1 systems capture the temporal rate and concentration of growth factors, and encode these distinct physical properties into transient and sustained ERK activation, respectively.

Dynamics of active Ras, active Rap1 and phosphorylated ERK were correctly reproduced with CellDesigner 3.0

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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

Name microMole

Definition µmol

# 2.2 Unit volume

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

Definition 1

#### 2.3 Unit area

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

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

# 2.4 Unit length

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

**Definition** m

# 2.5 Unit time

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

**Definition** s

# 3 Compartments

This model contains two compartments.

Table 2: Properties of all compartments.

| Id             | Name | SBO | Spatial Dimensions | Size | Unit           | Constant | Outside     |
|----------------|------|-----|--------------------|------|----------------|----------|-------------|
| compartment c1 |      |     | 3 3                | 1 1  | litre<br>litre | <b>✓</b> | compartment |

# 3.1 Compartment compartment

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

# 3.2 Compartment c1

This is a three dimensional compartment with a constant size of one litre, which is surrounded by compartment.

# 4 Species

This model contains 99 species. The boundary condition of five of these species is set to true so that these species' amount cannot be changed by any reaction. Section 6 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<br>Condi- |
|-------------------|------|-------------|------------------------|----------|--------------------|
|                   |      |             |                        |          | tion               |
| EGFR              |      | compartment | $\mu mol \cdot l^{-1}$ |          |                    |
| L_EGFR            |      | compartment | $\mu mol \cdot l^{-1}$ |          |                    |
| $L\_EGFR\_dimer$  |      | compartment | $\mu mol \cdot l^{-1}$ |          |                    |
| SOS               |      | c1          | $\mu mol \cdot l^{-1}$ |          |                    |
| $L_dpEGFR$        |      | compartment | $\mu mol \cdot l^{-1}$ | $\Box$   |                    |
| pSOS              |      | c1          | $\mu mol \cdot l^{-1}$ | $\Box$   |                    |
| SOS_Grb2          |      | c1          | $\mu mol \cdot l^{-1}$ | $\Box$   |                    |
| Grb2              |      | c1          | $\mu mol \cdot l^{-1}$ | $\Box$   |                    |
| Dok               |      | c1          | $\mu mol \cdot l^{-1}$ | $\Box$   |                    |
| pDok              |      | c1          | $\mu mol \cdot l^{-1}$ | $\Box$   |                    |
| Crk               |      | c1          | $\mu mol \cdot l^{-1}$ |          |                    |
| FRS2              |      | c1          | $\mu mol \cdot l^{-1}$ |          |                    |
| Shc               |      | c1          | $\mu mol \cdot l^{-1}$ |          |                    |
| pSOS_Grb2         |      | c1          | $\mu mol \cdot l^{-1}$ | $\Box$   |                    |
| Rap1_GDP          |      | c1          | $\mu mol \cdot l^{-1}$ | $\Box$   |                    |
| MEK               |      | c1          | $\mu mol \cdot l^{-1}$ | $\Box$   |                    |
| MKP3              |      | c1          | $\mu mol \cdot l^{-1}$ |          |                    |
| $pShc\_dpEGFR$    |      | c1          | $\mu mol \cdot l^{-1}$ |          |                    |
| dpEGFR_c_Cbl      |      | c1          | $\mu mol \cdot l^{-1}$ |          |                    |
| B_Raf_Rap1_GTP    |      | c1          | $\mu mol \cdot l^{-1}$ | $\Box$   |                    |
| pShc_dpEGFR_c_Cbl |      | c1          | $\mu mol \cdot l^{-1}$ |          | $\Box$             |

| Id                          | Name | Compartment | Derived Unit                             | Constant | Boundary<br>Condi-<br>tion |
|-----------------------------|------|-------------|--|----------|----------------------------|
| pFRS2_dpEGFR_c_Cbl          |      | c1          | $\mu$ mol· $1^{-1}$                      |          | $\Box$                     |
| Shc_dpEGFR                  |      | c1          | $\mu mol \cdot l^{-1}$                   | $\Box$   | $\Box$                     |
| c_Cbl                       |      | c1          | $\mu mol \cdot l^{-1}$                   | $\Box$   | $\Box$                     |
| RasGAP                      |      | c1          | $\mu mol \cdot l^{-1}$                   | $\Box$   | $\Box$                     |
| c_Raf                       |      | c1          | $\mu mol \cdot l^{-1}$                   | $\Box$   | $\Box$                     |
| B_Raf                       |      | c1          | $\mu$ mol $\cdot$ l $^{-1}$              | $\Box$   | $\Box$                     |
| ERK                         |      | c1          | $\mu$ mol $\cdot$ l $^{-1}$              | $\Box$   | $\Box$                     |
| PP2A                        |      | c1          | $\mu$ mol $\cdot$ l $^{-1}$              | $\Box$   | $\Box$                     |
| Ras_GDP                     |      | c1          | $\mu$ mol $\cdot$ l $^{-1}$              | $\Box$   | $\Box$                     |
| Rap1GAP                     |      | c1          | $\mu mol \cdot l^{-1}$                   | $\Box$   | $\Box$                     |
| C3G                         |      | c1          | $\mu \mathrm{mol} \cdot \mathrm{l}^{-1}$ | $\Box$   | $\Box$                     |
| NGFR                        |      | compartment | $\mu \mathrm{mol} \cdot \mathrm{l}^{-1}$ | $\Box$   | $\Box$                     |
| pShc                        |      | c1          | $\mu$ mol $\cdot$ l $^{-1}$              | $\Box$   | $\Box$                     |
| pFRS2_dpEGFR                |      | c1          | $\mu mol \cdot l^{-1}$                   | $\Box$   | $\Box$                     |
| pTrkA_endo                  |      | c1          | $\mu$ mol $\cdot$ l $^{-1}$              |          | $\Box$                     |
| MEK_ERK                     |      | c1          | $\mu$ mol $\cdot$ l $^{-1}$              |          | $\Box$                     |
| pMEK_ERK                    |      | c1          | $\mu \mathrm{mol} \cdot \mathrm{l}^{-1}$ |          | $\Box$                     |
| FRS2_dpEGFR_c_Cbl-          |      | c1          | $\mu mol \cdot l^{-1}$                   | $\Box$   | $\Box$                     |
| $\_{\tt ubiq}$              |      |             |  |          |                            |
| Crk_C3G_pFRS2-              |      | c1          | $\mu$ mol $\cdot$ l $^{-1}$              |          | $\Box$                     |
| $\_\mathtt{dpEGFR\_c\_Cbl}$ |      |             |  |          |                            |
| pShc_dpEGFR_c_Cbl-          |      | c1          | $\mu$ mol $\cdot$ l $^{-1}$              |          | $\Box$                     |
| _ubiq                       |      |             |  |          |                            |
| Crk_C3G_pFRS2-              |      | c1          | $\mu mol \cdot l^{-1}$                   |          |                            |
| $_{	t dpEGFR}$              |      |             |  |          |                            |
| Grb2_SOS_pShc-              |      | c1          | $\mu mol \cdot l^{-1}$                   | $\Box$   | $\Box$                     |
| $\_dpEGFR\_c\_Cbl\_ubiq$    |      |             |  |          |                            |

| Id                        | Name       | Compartment | Derived Unit              | Constant | Boundary<br>Condi-<br>tion |
|---------------------------|------------|-------------|---------------------------|----------|----------------------------|
| Grb2_SOS_pShc-            |            | c1          | $\mu$ mol·l <sup>-1</sup> |          |                            |
| _dpEGFR_c_Cbl             |            |             |                           |          |                            |
| Shc_dpEGFR_c_Cbl-         |            | c1          | $\mu mol \cdot l^{-1}$    |          | $\Box$                     |
| _ubiq                     |            |             |                           |          |                            |
| dpEGFR_c_Cbl_ubiq         |            | c1          | $\mu mol \cdot l^{-1}$    |          | $\Box$                     |
| proteosome                | proteasome | c1          | $\mu mol \cdot l^{-1}$    |          | $\Box$                     |
| Grb2_SOS_pShc             |            | c1          | $\mu mol \cdot l^{-1}$    |          | $\Box$                     |
| Shc_dpEGFR_c_Cbl          |            | c1          | $\mu mol \cdot l^{-1}$    |          | $\Box$                     |
| Grb2_SOS_pShc-<br>_dpEGFR |            | c1          | $\mu mol \cdot l^{-1}$    |          |                            |
| pFRS2                     |            | c1          | $\mu$ mol·l <sup>-1</sup> |          |                            |
| FRS2_dpEGFR               |            | c1          | $\mu$ mol·l <sup>-1</sup> |          |                            |
| pDok_RasGAP               |            | c1          | $\mu$ mol·l <sup>-1</sup> |          |                            |
| pMEK                      |            | c1          | $\mu$ mol·l <sup>-1</sup> |          | $\Box$                     |
| FRS2_dpEGFR_c_Cbl         |            | c1          | $\mu$ mol·l <sup>-1</sup> |          | $\Box$                     |
| pFRS2_dpEGFR_c-           |            | c1          | $\mu mol \cdot l^{-1}$    |          | $\Box$                     |
| _Cbl_ubiq                 |            |             |                           |          |                            |
| Ras_GTP                   |            | c1          | $\mu mol \cdot l^{-1}$    |          | $\Box$                     |
| Crk_C3G_pFRS2-            |            | c1          | $\mu mol \cdot l^{-1}$    |          | $\Box$                     |
| _dpEGFR_c_Cbl_ubiq        |            |             |                           |          |                            |
| c_Raf_Ras_GTP             |            | c1          | $\mu$ mol·l <sup>-1</sup> |          | $\Box$                     |
| B_Raf_Ras_GTP             |            | c1          | $\mu$ mol·l <sup>-1</sup> |          | $\Box$                     |
| ррМЕК                     |            | c1          | $\mu$ mol·l <sup>-1</sup> |          | $\Box$                     |
| ppERK                     |            | c1          | $\mu$ mol·l <sup>-1</sup> |          | $\Box$                     |
| pTrkA                     |            | compartment | $\mu$ mol·l <sup>-1</sup> |          |                            |
| Crk_C3G                   |            | c1          | $\mu$ mol·l <sup>-1</sup> |          |                            |
| Rap1_GTP                  |            | c1          | $\mu mol \cdot l^{-1}$    |          |                            |

| Id                  | Name           | Compartment | Derived Unit                             | Constant | Boundary<br>Condi-<br>tion |
|---------------------|----------------|-------------|--|----------|----------------------------|
| $_{\rm L\_NGFR}$    |                | compartment | $\mu mol \cdot l^{-1}$                   |          |                            |
| ppMEK_ERK           |                | c1          | $\mu$ mol·l <sup>-1</sup>                |          | $\Box$                     |
| dppERK              |                | c1          | $\mu mol \cdot l^{-1}$                   |          | $\Box$                     |
| Shc_pTrkA           |                | c1          | $\mu \mathrm{mol} \cdot \mathrm{l}^{-1}$ | $\Box$   | $\Box$                     |
| Shc_pTrkA_endo      |                | c1          | $\mu \mathrm{mol} \cdot \mathrm{l}^{-1}$ | $\Box$   | $\Box$                     |
| pShc_pTrkA          |                | c1          | $\mu mol \cdot l^{-1}$                   |          | $\Box$                     |
| pFRS2_pTrkA         |                | c1          | $\mu mol \cdot l^{-1}$                   |          |                            |
| FRS2_pTrkA          |                | c1          | $\mu mol \cdot l^{-1}$                   |          |                            |
| pShc_pTrkA_endo     |                | c1          | $\mu mol \cdot l^{-1}$                   |          |                            |
| FRS2_pTrkA_endo     |                | c1          | $\mu mol \cdot l^{-1}$                   |          | $\Box$                     |
| pFRS2_pTrkA_endo    |                | c1          | $\mu mol \cdot l^{-1}$                   |          | $\Box$                     |
| Crk_C3G_pFRS2-      |                | c1          | $\mu mol \cdot l^{-1}$                   |          | $\Box$                     |
| $_{	t pTrkA\_endo}$ |                |             |  |          |                            |
| Grb2_SOS_pShc-      |                | c1          | $\mu mol \cdot l^{-1}$                   |          |                            |
| _pTrkA              |                |             |  |          |                            |
| Crk_C3G_pFRS2-      |                | c1          | $\mu$ mol·l <sup>-1</sup>                |          | $\Box$                     |
| _pTrkA              |                |             |  |          |                            |
| Grb2_SOS_pShc-      |                | c1          | $\mu mol \cdot l^{-1}$                   |          |                            |
| _pTrkA_endo         |                |             |  |          |                            |
| c_Raf_Ras_GTP_MEK   |                | c1          | $\mu mol \cdot l^{-1}$                   |          |                            |
| c_Raf_Ras_GTP_pME   | K.K            | c1          | $\mu mol \cdot l^{-1}$                   |          | $\Box$                     |
| c_Raf_Ras_GTP_MEK   | ; <del>-</del> | c1          | $\mu mol \cdot l^{-1}$                   | $\Box$   | $\Box$                     |
| _ERK                |                |             |  |          |                            |
| c_Raf_Ras_GTP-      |                | c1          | $\mu mol \cdot l^{-1}$                   | $\Box$   | $\Box$                     |
| _pMEK_ERK           |                |             |  |          |                            |
| B_Raf_Ras_GTP_MEK   |                | c1          | $\mu mol \cdot l^{-1}$                   |          |                            |
| $B_Raf_Ras_GTP_pME$ | K.K            | c1          | $\mu mol \cdot l^{-1}$                   | $\Box$   |                            |

| Id           | Name    | Compartment | Derived Unit                             | Constant | Boundary<br>Condi-<br>tion |
|--------------|---------|-------------|--|----------|----------------------------|
| B_Raf_Ras_G7 | TP_MEK- | c1          | $\mu mol \cdot l^{-1}$                   | $\Box$   |                            |
| _ERK         |         |             |  |          |                            |
| B_Raf_Ras_G7 | P-      | c1          | $\mu mol \cdot l^{-1}$                   |          | $\Box$                     |
| _pMEK_ERK    |         |             |  |          |                            |
| B_Raf_Rap1_0 | TTP_MEK | c1          | $\mu$ mol·l <sup>-1</sup>                |          |                            |
| B_Raf_Rap1_0 | TP-     | c1          | $\mu$ mol·l <sup>-1</sup>                |          |                            |
| _pMEK        |         |             |  |          |                            |
| B_Raf_Rap1_0 | STP-    | c1          | $\mu$ mol·l <sup>-1</sup>                |          | $\Box$                     |
| _MEK_ERK     |         |             |  |          |                            |
| B_Raf_Rap1_0 | STP-    | c1          | $\mu$ mol·l <sup>-1</sup>                |          |                            |
| _pMEK_ERK    |         |             |  |          |                            |
| ppERK_MKP3   |         | c1          | $\mu \mathrm{mol} \cdot \mathrm{l}^{-1}$ |          |                            |
| dppERK_MKP3  |         | c1          | $\mu$ mol·l <sup>-1</sup>                |          |                            |
| pro_TrkA     |         | c1          | $\mu mol \cdot l^{-1}$                   |          |                            |
| NGF          |         | compartment | $\mu mol \cdot l^{-1}$                   |          |                            |
| EGF          |         | compartment | $\mu mol \cdot l^{-1}$                   |          |                            |
| pro_EGFR     |         | c1          | $\mu$ mol·l <sup>-1</sup>                |          |                            |
| degradation  | ı       | c1          | $\mu \mathrm{mol} \cdot \mathrm{l}^{-1}$ |          |                            |

# **5 Reactions**

This model contains 150 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 4: Overview of all reactions

| $N_{\bar{0}}$ | Id  | Name                               | Reaction Equation   | SBO |
|---------------|-----|------------------------------------|---|-----|
| 1             | re1 | form_EGFreceptor                   | pro_EGFR <del>←</del> EGFR  |     |
| 2             | re2 | EGFbinding                         | $EGF + EGFR \Longrightarrow L\_EGFR$                                    |     |
| 3             | re8 | dimerization                       | $2 L\_EGFR \Longrightarrow L\_EGFR\_dimer$                              |     |
| 4             | J3  | binding_SOS_Grb2                   | $SOS + Grb2 \Longrightarrow SOS\_Grb2$                                  |     |
| 5             | J4  | binding_pSOS_Grb2                  | $Grb2 + pSOS \Longrightarrow pSOS\_Grb2$                                |     |
| 6             | J5  | EGFRphosphorylation                | $L\_EGFR\_dimer \Longrightarrow L\_dpEGFR$                              |     |
| 7             | J6  | binding_cCbI_dpEGFR                | $L_dpEGFR + c_Cbl \Longrightarrow dpEGFR_c_Cbl$                         |     |
| 8             | J7  | binding_pShc_LdpEGFR               | $L_dpEGFR + pShc \Longrightarrow pShc_dpEGFR$                           |     |
| 9             | Ј8  | pDOKdephosphorylation              | $pDok \rightleftharpoons Dok$   |     |
| 10            | J9  | binding_cCbl_pShc_dpEGFR           | $c_Cbl + pShc_dpEGFR \Longrightarrow pShc_dpEGFR_c_Cbl$                 |     |
| 11            | J10 | SOSdephosphorylation               | $pSOS \longrightarrow SOS$  |     |
| 12            | J11 | pSOS_Grb2_dephosphorylation        | $pSOS\_Grb2 \longrightarrow SOS\_Grb2$                                  |     |
| 13            | J12 | binding_Shc_LdpEGFR                | $L_dpEGFR + Shc \Longrightarrow Shc_dpEGFR$                             |     |
| 14            | J13 | Shc_dpEGFR_phosphorylation         | $Shc\_dpEGFR \longrightarrow pShc\_dpEGFR$                              |     |
| 15            | J14 | dpEGFR_c_Cbl_ubiquitination        | dpEGFR_c_Cbl dpEGFR_c_Cbl_ubiq  |     |
| 16            | J15 | dpEGFR_cCbl_degrad                 | $dpEGFR_c\_Cbl\_ubiq \longrightarrow proteosome + c\_Cbl$               |     |
| 17            | J16 | binding_cCbl_Shc_dpEGFR            | $c_Cbl + Shc_dpEGFR \Longrightarrow Shc_dpEGFR_c_Cbl$                   |     |
| 18            | J17 | Shc_dpEGFR_c_CBl_Ubiquitination    | Shc_dpEGFR_c_Cbl> Shc_dpEGFR_c_Cbl_ubiq                                 |     |
| 19            | J18 | Shc_dpEGFR_c_Cbl_ubiq_Degradation  | $Shc\_dpEGFR\_c\_Cbl\_ubiq \longrightarrow proteosome + c\_Cbl + Shc$   |     |
| 20            | J19 | pShc_dpEGFR_c_Cbl_ubiquitination   | pShc_dpEGFR_c_Cbl pShc_dpEGFR_c_Cbl_ubi                                 | q   |
| 21            | J20 | pShc_dpEGFR_c_Cbl_ubiq_degradation | $pShc\_dpEGFR\_c\_Cbl\_ubiq \longrightarrow proteosome + c\_Cbl + pShc$ |     |

\_ubiquitination

| N⁰       | Id  | Name                                 | Reaction Equation                                      | SBO             |
|----------|-----|--------------------------------------|--|-----------------|
| 46       | J46 | FRS2_dpEGFR_c_Cbl_ubiq_dissociation  | FRS2_dpEGFR_c_Cbl_ubiq> proteosome                     | +               |
|          |     | •                                    | c_Cbl+FRS2   |                 |
| 47       | J47 | pFRS2_dpEGFR_c_Cbl_ubiq_dissociation | pFRS2_dpEGFR_c_Cbl_ubiq> proteosome                    | +               |
|          |     |                                      | c_Cbl+pFRS2  |                 |
| 48       | J49 | binding_RasGAP_to_pDOK               | $pDok + RasGAP \Longrightarrow pDok\_RasGAP$           |                 |
| 49       | J50 | SOS_Grb2_phosphorylation             | $SOS\_Grb2 \xrightarrow{dppERK} pSOS\_Grb2$            |                 |
|          | 300 |                                      |  |                 |
| 50       | J51 | SOS_phosphorylation                  | $SOS \xrightarrow{dppERK} pSOS$                        |                 |
| 51       | J52 | binding_c_Raf_to_Ras_GTP             | $c_Raf + Ras_GTP \Longrightarrow c_Raf_Ras_GTP$        |                 |
| 52       | J53 | binding_B_Raf_to_Rap1_GTP            | $Rap1\_GTP + B\_Raf \Longrightarrow B\_Raf\_Rap1\_GTP$ |                 |
| 53       | J54 | binding_B_Raf_to_Ras_GTP             | $Ras\_GTP + B\_Raf \Longrightarrow B\_Raf\_Ras\_GTP$   |                 |
| 54       | J57 | ppMEK_dephosphorylation              | $ppMEK \xrightarrow{PP2A} pMEK$                        |                 |
| 55       | J58 | pMEK_dephosphorylation               | $pMEK \xrightarrow{PP2A} MEK$                          |                 |
| 56       | J61 | ppMEK_ERK                            | $ppMEK\_ERK \xrightarrow{PP2A} pMEK\_ERK$              |                 |
| 57       | J62 | pMEK_ERK_dephosphorylation           | pMEK_ERK $\xrightarrow{PP2A}$ MEK_ERK                  |                 |
| 58       | J63 | ppERK_dimerization                   | 2 ppERK ⇒ dppERK                                       |                 |
| 59       | J66 | Ras_GTP_dephosphorylation            | $Ras\_GTP \longrightarrow Ras\_GDP$                    |                 |
| 60       | J67 | Rap1_GTP_dephosphorylation           | $Rap1\_GTP \longrightarrow Rap1\_GDP$                  |                 |
| 61       | J68 | Rap1_GTP_phosphorylation             | Rap1_GDP Crk_C3G_pFRS2_dpEGFR_c_Cbl, (                 | Crk_C3G_pFRS2_d |
| 62       | J69 | Ras_GDP_phosphorylation              | Ras_GDP Grb2_SOS_pShc_dpEGFR_c_Cbl, Gr                 | b2_SOS_pShc_dpE |
| 63       | J70 | binding_NGF_to_NGFR                  | $NGF + NGFR \Longrightarrow L\_NGFR$                   |                 |
| 64       | J71 | TrkA_phosphorylation                 | $L_{NGFR} \longrightarrow pTrkA$                       |                 |
| 65       | J72 | pTrkA_intermalization                | •  |                 |
|          |     | •                                    | pTrkA → pTrkA_endo                                     |                 |
| 66<br>67 | J73 | pTrkA_endo_degradation               | pTrkA_endo → degradation                               |                 |
| 67<br>68 | J74 | pTrkA_degradation                    | pTrkA → degradation                                    |                 |
| Uð       | J75 | binding_Shc_to_pTrkA                 | $Shc + pTrkA \Longrightarrow Shc_pTrkA$                |                 |

| 12                           | Nº Id   | Name                                | Reaction Equation   | SBO          |
|------------------------------|---------|-------------------------------------|---|--------------|
|                              | 69 J76  | binding_pShc_to_pTrkA               | $pShc + pTrkA \Longrightarrow pShc_pTrkA$                                   |              |
|                              | 70 J77  | binding_FRS2_to_pTrkA               | $FRS2 + pTrkA \Longrightarrow FRS2 pTrkA$                                   |              |
|                              | 71 J78  | binding_pFRS2_to_pTrkA              | $pFRS2 + pTrkA \Longrightarrow pFRS2 pTrkA$                                 |              |
|                              | 72 J79  | binding_Shc_to_pTrkA_endo           | $pTrkA\_endo + Shc \Longrightarrow Shc\_pTrkA\_endo$                        |              |
|                              | 73 J80  | binding_pShc_to_pTrkA_endo          | $pTrkA\_endo + pShc \Longrightarrow pShc\_pTrkA\_endo$                      |              |
|                              | 74 J81  | Shc_pTrkA_endo_phosphorylation      | $Shc_pTrkA_endo \longrightarrow pShc_pTrkA_endo$                            |              |
|                              | 75 J82  | Shc_pTrkA_phosphorylation           | $Shc_pTrkA \longrightarrow pShc_pTrkA$                                      |              |
|                              | 76 J83  | pFRS2_pTrkA_phosphorylation         | $FRS2_pTrkA \longrightarrow pFRS2_pTrkA$                                    |              |
|                              | 77 J84  | binding_FRS2_to_pTrkA_endo          | $pTrkA\_endo + FRS2 \Longrightarrow FRS2\_pTrkA\_endo$                      |              |
|                              | 78 J85  | binding_pFRS2_to_pTrkA_endo         | $pTrkA\_endo + pFRS2 \Longrightarrow pFRS2\_pTrkA\_endo$                    |              |
| $^{p}$ ro                    | 79 J86  | FRS2_pTrkA_endo_phosphorylation     | $FRS2\_pTrkA\_endo \longrightarrow pFRS2\_pTrkA\_endo$                      |              |
| duc                          | 80 J87  | FRS2_pTrkA_degradation              | $FRS2_pTrkA \longrightarrow degradation + FRS2$                             |              |
| Produced by SBML2PTEX        | 81 J88  | pFRS2_pTrkA_degradation             | $pFRS2\_pTrkA \longrightarrow degradation + pFRS2$                          |              |
| by                           | 82 J89  | Shc_pTrkA_degradation               | $Shc_pTrkA \longrightarrow degradation + Shc$                               |              |
| <u>88</u>                    | 83 J90  | pShc_pTrkA_degradation              | $pShc_pTrkA \longrightarrow degradation + pShc$                             |              |
| $\leq$                       | 84 J92  | FRS2_pTrkA_endo_degradation         | $FRS2\_pTrkA\_endo \longrightarrow degradation + FRS2$                      |              |
| Ä                            | 85 J93  | Shc_pTrkA_endo_degradation          | $Shc_pTrkA_endo \longrightarrow degradation + Shc$                          |              |
| $\stackrel{\square}{\times}$ | 86 J94  | pShc_pTrkA_endo_degradation         | $pShc_pTrkA_endo \longrightarrow degradation + pShc$                        |              |
|                              | 87 J95  | binding_Grb2_SOS_to_pShc_pTrkA_endo | $SOS\_Grb2+pShc\_pTrkA\_endo \Longrightarrow Grb2\_SOS\_pShc$               | _pTrkA_endo  |
|                              | 88 J96  | binding_Grb2_SOS_to_pShc_pTrkA      | $SOS\_Grb2+pShc\_pTrkA \Longrightarrow Grb2\_SOS\_pShc\_pTrk$               | Α            |
|                              | 89 J97  | Grb2_SOS_pShc_pTrkA_ubiquitination  | $Grb2\_SOS\_pShc\_pTrkA \longrightarrow Grb2\_SOS\_pShc\_pTrkA$             | _endo        |
|                              | 90 J98  | Crk_C3G_pFRS2_pTrkA_ubiquitination  | $Crk_{-}C3G_{-}pFRS2_{-}pTrkA \longrightarrow Crk_{-}C3G_{-}pFRS2_{-}pTrkA$ | A_endo       |
|                              | 91 J99  | pFRS2_pTrkA_ubiquitination          | $pFRS2\_pTrkA \longrightarrow pFRS2\_pTrkA\_endo$                           |              |
|                              | 92 J100 | FRS2_pTrkA_ubiquitination           | $FRS2_pTrkA \longrightarrow FRS2_pTrkA\_endo$                               |              |
|                              | 93 J101 | pShc_pTrkA_ubiquitination           | $pShc\_pTrkA \longrightarrow pShc\_pTrkA\_endo$                             |              |
|                              | 94 J102 | Shc_pTrkA_ubiquitination            | $Shc_pTrkA \longrightarrow Shc_pTrkA_endo$                                  |              |
|                              | 95 J103 | binding_Crk_C3G_to_pFRS2_pTrkA      | $Crk\_C3G+pFRS2\_pTrkA \Longrightarrow Crk\_C3G\_pFRS2\_pTr$                | ·kA          |
|                              | 96 J104 | binding_Crk_C3G_to_pFRS2_pTrkA_endo | $Crk_C3G + pFRS2_pTrkA_endo \rightleftharpoons Crk_C3G_pFRS$                | 2_pTrkA_endo |

| N⁰  | Id   | Name  | Reaction Equation  | SBO                            |
|-----|------|---|--|--------------------------------|
| 97  | J105 | binding_Grb2_SOS_pShc_to_pTrkA              | Grb2_SOS_pShc  | +                              |
|     |      |   | pTrkA ⇒ Grb2_SOS_pShc_pTrkA                                  |                                |
| 98  | J106 | binding_Grb2_SOS_pShc_to_pTrkA_endo         | Grb2_SOS_pShc  | +                              |
|     |      |   | pTrkA_endo \ Grb2_SOS_pShc_pTrkA_endo                        |                                |
| 99  | J107 | Crk_C3G_pFRS2_pTrkA_degradation             | $Crk_C3G_pFRS2_pTrkA \longrightarrow degradation$            | +                              |
|     |      |   | pFRS2+Crk_C3G  |                                |
| 100 | J108 | $Crk\_C3G\_pFRS2\_pTrkA\_endo\_degradation$ | $Crk\_C3G\_pFRS2\_pTrkA\_endo \longrightarrow degradation$   | +                              |
|     |      |   | $Crk\_C3G + pFRS2$   |                                |
| 101 | J109 | Grb2_SOS_pShc_pTrkA_degradation             | $Grb2\_SOS\_pShc\_pTrkA \longrightarrow degradation$         | +                              |
|     |      |   | Grb2_SOS_pShc  |                                |
| 102 | J110 | Grb2_SOS_pShc_pTrkA_endo_degradation        | $Grb2\_SOS\_pShc\_pTrkA\_endo \longrightarrow degradation$   | +                              |
|     |      |   | Grb2_SOS_pShc  |                                |
| 103 | J112 | pFRS2_pTrkA_endo_degradation                | $pFRS2\_pTrkA\_endo \longrightarrow degradation + pFRS2$     |                                |
| 104 | J113 | form_NGFreceptor                            | $pro\_TrkA \Longrightarrow NGFR$                             |                                |
| 105 | J115 | binding_Shc_to_dpEGFR_c_Cbl                 | $Shc + dpEGFR\_c\_Cbl \Longrightarrow Shc\_dpEGFR\_c\_Cbl$   |                                |
| 106 | J116 | binding_pShc_to_dpEGFR_c_Cbl                | $dpEGFR\_c\_Cbl + pShc \Longrightarrow pShc\_dpEGFR\_c\_Cbl$ | 1                              |
| 107 | J117 | binding_SOS_Grb2_to_pShc_dpEGFR_c_Cbl       | pShc_dpEGFR_c_Cbl  | +                              |
|     |      |   | $SOS\_Grb2 \Longrightarrow Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl$  |                                |
| 108 | J119 | binding_c_Cbl_to_Crk_C3G_pFRS2_dpEGFR       | $c\_Cbl+Crk\_C3G\_pFRS2\_dpEGFR \Longrightarrow Crk\_C3G$    | _pFRS2_dpEGFR_c_Cbl            |
| 109 | J118 | binding_FRS2_to_dpEGFR_c_Cbl                | $dpEGFR_c\_Cbl + FRS2 \Longrightarrow FRS2\_dpEGFR_c\_Cbl$   |                                |
| 110 | J120 | binding_pFRS2_to_dpEGFR_c_Cbl               | $dpEGFR\_c\_Cbl+pFRS2 \Longrightarrow pFRS2\_dpEGFR\_c\_c$   | .Cbl                           |
| 111 | J121 | Ras_GTP_dephosphorylation                   | $Ras\_GTP \xrightarrow{pDok\_RasGAP} Ras\_GDP$               |                                |
| 112 | J122 | RAP1_GTP_dephosphorylation                  | $Rap1\_GTP \xrightarrow{Rap1GAP} Rap1\_GDP$                  |                                |
| 113 | J123 | Dok_phosphorylation                         | Dok PTrkA, Shc_pTrkA, pShc_pTrkA, Grb2_SOS                   | _pShc_pTrkA, FRS2_pTrkA, pFRS2 |
| 114 | J124 | Grb1_SOS_pShc_dissociation                  | $Grb2\_SOS\_pShc \xrightarrow{dppERK} pShc + pSOS\_Grb2$     |                                |
| 115 | J133 | binding_MEK_to_ERK                          | $ERK + MEK \Longrightarrow MEK\_ERK$                         |                                |
| 116 | J134 | binding_ERK_to_pMEK                         | $ERK + pMEK \Longrightarrow pMEK\_ERK$                       |                                |
|     |      |   |  |                                |

| 14                    | N⁰  | Id      | Name                        | Reaction Equation  | SBO       |
|-----------------------|-----|---------|-----------------------------|--|-----------|
|                       | 117 | J135    | binding_ERK_to_ppMEK        | $ERK + ppMEK \Longrightarrow ppMEK\_ERK$                               |           |
|                       | 118 | J136    | ppMEK_ERK_dissociation      | $ppMEK\_ERK \longrightarrow ppERK + ppMEK$                             |           |
|                       | 119 | J137    | c_Raf_Ras_GTP_dissociation  | $c_Raf_Ras_GTP \xrightarrow{pDok_RasGAP} c_Raf_Ras_GDP$                |           |
|                       | 120 | J138    | B_Raf_Ras_GTP_dissociation  | $B\_Raf\_Ras\_GTP \xrightarrow{pDok\_RasGAP} B\_Raf + Ras\_GDP$        |           |
|                       | 121 | J139    | B_Raf_Rap1_GTP_dissociation | $B_Raf_Rap1_GTP \xrightarrow{Rap1GAP} B_Raf_Rap1_GDP$                  |           |
|                       | 122 | J140    |                             | $c_Raf_Ras_GTP + MEK \longrightarrow c_Raf_Ras_GTP_MEK$                |           |
|                       | 123 | J141    |                             | c_Raf_Ras_GTP+pMEK === c_Raf_Ras_GTP_pMEk                              | ζ         |
|                       | 124 | J142    |                             | c_Raf_Ras_GTP+MEK_ERK === c_Raf_Ras_GTP_M                              | 1EK_ERK   |
| P                     | 125 | J143    |                             | c_Raf_Ras_GTP+pMEK_ERK \equiv c_Raf_Ras_GTP_                           | pMEK_ERK  |
| rod                   | 126 | J144    |                             | B_Raf_Ras_GTP+MEK 	⇒ B_Raf_Ras_GTP_MEK                                 | •         |
| исє                   | 127 | J145    |                             | B_Raf_Ras_GTP+pMEK === B_Raf_Ras_GTP_pME                               | K         |
| l be                  | 128 | J146    |                             | B_Raf_Ras_GTP+MEK_ERK \Rightarrow B_Raf_Ras_GTP_                       | MEK_ERK   |
| )y (                  | 129 | J147    |                             | B_Raf_Ras_GTP+pMEK_ERK <del>====================================</del> | _pMEK_ERK |
| Produced by SBML2PTEX | 130 | J148    |                             | B_Raf_Rap1_GTP +   | •         |
| <u> </u>              |     |         |                             | MEK <del>← B_Raf_Rap1_GTP_MEK</del>                                    |           |
| A                     | 131 | J149    |                             | B_Raf_Rap1_GTP +   |           |
| '×'                   |     |         |                             | pMEK ⇒ B_Raf_Rap1_GTP_pMEK   |           |
|                       | 132 | J150    |                             | B_Raf_Rap1_GTP +   |           |
|                       |     |         |                             | MEK_ERK ⇒ B_Raf_Rap1_GTP_MEK_ERK                                       |           |
|                       | 133 | J151    |                             | B_Raf_Rap1_GTP +   |           |
|                       |     |         |                             | pMEK_ERK \=== B_Raf_Rap1_GTP_pMEK_ERK                                  |           |
|                       | 134 | J152    |                             | $c_Raf_Ras_GTP_MEK \longrightarrow c_Raf_Ras_GTP$ +                    |           |
|                       |     |         |                             | pMEK   |           |
|                       | 135 | J153    |                             | $c_Raf_Ras_GTP_pMEK \longrightarrow c_Raf_Ras_GTP +$                   |           |
|                       |     |         |                             | ррМЕК  |           |
|                       | 136 | J154    |                             | c_Raf_Ras_GTP_MEK_ERK → c_Raf_Ras_GTP +                                |           |
|                       | 120 | <b></b> |                             | pMEK_ERK   |           |
|                       |     |         |                             | r  |           |

| N⁰    | Id           | Name | Reaction Equation SBO  |  |
|-------|--------------|------|--|--|
| 137   | J155         |      | c_Raf_Ras_GTP_pMEK_ERK → c_Raf_Ras_GTP+  |  |
|       |              |      | ppMEK_ERK  |  |
| 138   | J156         |      | $B_Raf_Ras_GTP\_MEK \longrightarrow B_Raf_Ras\_GTP +$                                      |  |
|       |              |      | pMEK   |  |
| 139   | J157         |      | $B_Raf_Ras_GTP_pMEK \longrightarrow B_Raf_Ras_GTP +$                                       |  |
|       |              |      | ppMEK  |  |
| 140   | J158         |      | $B_Raf_Ras_GTP_MEK_ERK \longrightarrow B_Raf_Ras_GTP +$                                    |  |
|       |              |      | pMEK_ERK   |  |
| 141   | J159         |      | $B_Raf_Ras_GTP_pMEK_ERK \longrightarrow B_Raf_Ras_GTP +$                                   |  |
|       |              |      | ppMEK_ERK  |  |
| 142   | J160         |      | $B_Raf_Rap1_GTP_MEK \longrightarrow B_Raf_Rap1_GTP +$                                      |  |
|       |              |      | pMEK   |  |
| 143   | J161         |      | $B_Raf_Rap1_GTP_pMEK \longrightarrow B_Raf_Rap1_GTP +$                                     |  |
| 1.1.1 |              |      | ppMEK  |  |
| 144   | J162         |      | B_Raf_Rap1_GTP_MEK_ERK → B_Raf_Rap1_GTP+   |  |
| 1.45  | 74.00        |      | pMEK_ERK   |  |
| 145   | J163         |      | B_Raf_Rap1_GTP_pMEK_ERK → B_Raf_Rap1_GTP+  |  |
| 1.46  | T1.C./       |      | ppMEK_ERK  |  |
| 146   | J164         |      | $Crk_C3G_pFRS2\_dpEGFR\_c\_Cbl\_ubiq \longrightarrow c\_Cbl + pFRS2 + Crk_C3G$             |  |
| 147   | J165         |      | •  |  |
| 147   | J165<br>J166 |      | $MKP3 + dppERK \longrightarrow dppERK\_MKP3$ $MKP3 + ppEPK \longrightarrow ppEPK\_MKP3$    |  |
| 148   |              |      | $MKP3 + ppERK \Longrightarrow ppERK\_MKP3$   |  |
|       | J167         |      | $ppERK\_MKP3 \longrightarrow ERK + MKP3$ $dpnERK\_MKP2 \longrightarrow ppERK + ERK + MKP2$ |  |
| 150   | J168         |      | $dppERK\_MKP3 \longrightarrow ppERK + ERK + MKP3$  |  |

#### **5.1 Reaction** re1

This is a reversible reaction of one reactant forming one product.

Name form\_EGFreceptor

# **Reaction equation**

$$pro\_EGFR \Longrightarrow EGFR$$
 (1)

#### Reactant

Table 5: Properties of each reactant.

| Id       | Name | SBO |
|----------|------|-----|
| pro_EGFR |      |     |

#### **Product**

Table 6: Properties of each product.

| Id   | Name | SBO |
|------|------|-----|
| EGFR |      |     |

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_1 = \text{vol} (\text{compartment}) \cdot (\text{rel\_k1} \cdot [\text{pro\_EGFR}] - \text{rel\_k2} \cdot [\text{EGFR}])$$
 (2)

Table 7: Properties of each parameter.

| Id     | Name | SBO | Value     | Unit | Constant       |
|--------|------|-----|-----------|------|----------------|
| re1_k1 |      |     | $10^{-4}$ |      | $\blacksquare$ |
| re1_k2 |      |     | $10^{-4}$ |      |                |

# 5.2 Reaction re2

This is a reversible reaction of two reactants forming one product.

Name EGFbinding

# **Reaction equation**

$$EGF + EGFR \Longrightarrow L EGFR \tag{3}$$

#### **Reactants**

Table 8: Properties of each reactant.

| Id   | Name | SBO |
|------|------|-----|
| EGF  |      |     |
| EGFR |      |     |

#### **Product**

Table 9: Properties of each product.

| Id     | Name | SBO |
|--------|------|-----|
| L_EGFR |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_2 = \text{vol}\left(\text{compartment}\right) \cdot \left(\text{re2\_k1} \cdot [\text{EGF}] \cdot [\text{EGFR}] - \text{re2\_k2} \cdot [\text{L\_EGFR}]\right)$$
 (4)

Table 10: Properties of each parameter.

| Id       | Name | SBO Value Unit | Constant       |
|----------|------|----------------|----------------|
| re2_k1   |      | 2.283          | $ \mathbf{Z} $ |
| $re2_k2$ |      | 0.003          | $\square$      |

#### 5.3 Reaction re8

This is a reversible reaction of one reactant forming one product.

Name dimerization

# **Reaction equation**

$$2L\_EGFR \Longrightarrow L\_EGFR\_dimer$$
 (5)

#### Reactant

Table 11: Properties of each reactant.

| Id     | Name | SBO |
|--------|------|-----|
| L_EGFR |      |     |

#### **Product**

Table 12: Properties of each product.

| Id           | Name | SBO |
|--------------|------|-----|
| L_EGFR_dimer |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_3 = \text{vol} (\text{compartment}) \cdot (\text{re8\_k1} \cdot [\text{L\_EGFR}] \cdot [\text{L\_EGFR}] - \text{re8\_k2} \cdot [\text{L\_EGFR\_dimer}])$$
 (6)

Table 13: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| re8_k1 |      | 10.00          |          |
| re8_k2 |      | 0.02           |          |

# **5.4 Reaction** J3

This is a reversible reaction of two reactants forming one product.

Name binding\_SOS\_Grb2

# **Reaction equation**

$$SOS + Grb2 \Longrightarrow SOS\_Grb2 \tag{7}$$

#### Reactants

Table 14: Properties of each reactant.

| Id   | Name | SBO |
|------|------|-----|
| SOS  |      |     |
| Grb2 |      |     |

#### **Product**

Table 15: Properties of each product.

| Id       | Name | SBO |
|----------|------|-----|
| SOS_Grb2 |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_4 = \text{vol}(c1) \cdot (J3\_k1 \cdot [SOS] \cdot [Grb2] - J3\_k2 \cdot [SOS\_Grb2])$$
(8)

Table 16: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J3_k1 |      | 0.030          | Ø        |
| J3_k2 |      | 0.017          |          |

# 5.5 Reaction J4

This is a reversible reaction of two reactants forming one product.

Name binding\_pSOS\_Grb2

# **Reaction equation**

$$Grb2 + pSOS \Longrightarrow pSOS\_Grb2$$
 (9)

#### **Reactants**

Table 17: Properties of each reactant.

| Id   | Name | SBO |
|------|------|-----|
| pSOS |      |     |

#### **Product**

Table 18: Properties of each product.

| Id        | Name | SBO |
|-----------|------|-----|
| pSOS_Grb2 |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_5 = \text{vol}(c1) \cdot (\text{J4}_k1 \cdot [\text{Grb2}] \cdot [\text{pSOS}] - \text{J4}_k2 \cdot [\text{pSOS}_G\text{rb2}])$$
 (10)

Table 19: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant       |
|---------|------|----------------|----------------|
| J4_k1   |      | 0.030          | $\overline{Z}$ |
| $J4_k2$ |      | 0.017          |                |

# **5.6 Reaction** J5

This is a reversible reaction of one reactant forming one product.

Name EGFRphosphorylation

# **Reaction equation**

$$L\_EGFR\_dimer \Longrightarrow L\_dpEGFR \tag{11}$$

#### Reactant

Table 20: Properties of each reactant.

| Id           | Name | SBO |
|--------------|------|-----|
| L_EGFR_dimer |      |     |

Table 21: Properties of each product.

| Id       | Name | SBO |
|----------|------|-----|
| L_dpEGFR |      |     |

**Derived unit** contains undeclared units

$$v_6 = vol\left(compartment\right) \cdot \left(J5\_k1 \cdot [L\_EGFR\_dimer] - J5\_k2 \cdot [L\_dpEGFR]\right) \tag{12}$$

Table 22: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J5_k1 |      | 4.000          |          |
| J5_k2 |      | 0.001          |          |

# **5.7 Reaction** J6

This is a reversible reaction of two reactants forming one product.

Name binding\_cCbI\_dpEGFR

# **Reaction equation**

$$L_dpEGFR + c_Cbl \Longrightarrow dpEGFR_c_Cbl$$
 (13)

#### **Reactants**

Table 23: Properties of each reactant.

| Id                | Name | SBO |
|-------------------|------|-----|
| L_dpEGFR<br>c_Cbl |      |     |

Table 24: Properties of each product.

| Id           | Name | SBO |
|--------------|------|-----|
| dpEGFR_c_Cbl |      |     |

**Derived unit** contains undeclared units

$$v_7 = \text{vol}(c1) \cdot (\text{J6\_k1} \cdot [\text{L\_dpEGFR}] \cdot [\text{c\_Cbl}] - \text{J6\_k2} \cdot [\text{dpEGFR\_c\_Cbl}])$$
 (14)

Table 25: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant |
|---------|------|----------------|----------|
| J6_k1   |      | 0.5            |          |
| $J6_k2$ |      | 0.2            |          |

#### 5.8 Reaction J7

This is a reversible reaction of two reactants forming one product.

Name binding\_pShc\_LdpEGFR

# **Reaction equation**

$$L_dpEGFR + pShc \Longrightarrow pShc_dpEGFR$$
 (15)

## **Reactants**

Table 26: Properties of each reactant.

| Id       | Name | SBO |
|----------|------|-----|
| L_dpEGFR |      |     |
| pShc     |      |     |

#### **Product**

Table 27: Properties of each product.

| Id          | Name | SBO |
|-------------|------|-----|
| pShc_dpEGFR |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_8 = \text{vol}(c1) \cdot (J7 k1 \cdot [L_dpEGFR] \cdot [pShc] - J7 k2 \cdot [pShc_dpEGFR])$$
 (16)

Table 28: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant     |
|-------|------|----------------|--------------|
| J7_k1 |      | 10.0           |              |
| J7_k2 |      | 0.2            | $\mathbf{Z}$ |

#### 5.9 Reaction J8

This is a reversible reaction of one reactant forming one product.

Name pDOKdephosphorylation

# **Reaction equation**

$$pDok \Longrightarrow Dok$$
 (17)

#### Reactant

Table 29: Properties of each reactant.

| Id   | Name | SBO |
|------|------|-----|
| pDok |      |     |

#### **Product**

Table 30: Properties of each product.

| Id  | Name | SBO |
|-----|------|-----|
| Dok |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_9 = \text{vol}(c1) \cdot (J8 \underline{k} \cdot [pDok] - J8 \underline{k} \cdot [Dok])$$

$$(18)$$

Table 31: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant       |
|-------|------|----------------|----------------|
| J8_k1 |      | 0.002          | $\overline{Z}$ |
| J8_k2 |      | $10^{-5}$      |                |

#### 5.10 Reaction J9

This is a reversible reaction of two reactants forming one product.

Name binding\_cCbl\_pShc\_dpEGFR

# **Reaction equation**

$$c\_Cbl + pShc\_dpEGFR \Longrightarrow pShc\_dpEGFR\_c\_Cbl$$
 (19)

#### **Reactants**

Table 32: Properties of each reactant.

| Id                   | Name | SBO |
|----------------------|------|-----|
| c_Cbl                |      |     |
| ${\tt pShc\_dpEGFR}$ |      |     |

#### **Product**

Table 33: Properties of each product.

| Id                | Name | SBO |
|-------------------|------|-----|
| pShc_dpEGFR_c_Cbl |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{10} = \text{vol}(c1) \cdot (\text{J9\_k1} \cdot [\text{c\_Cbl}] \cdot [\text{pShc\_dpEGFR}] - \text{J9\_k2} \cdot [\text{pShc\_dpEGFR\_c\_Cbl}]) \quad (20)$$

Table 34: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J9_k1 |      | 0.5            |          |
| J9_k2 |      | 0.2            |          |

#### 5.11 Reaction J10

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

Name SOSdephosphorylation

# **Reaction equation**

$$pSOS \longrightarrow SOS \tag{21}$$

#### Reactant

Table 35: Properties of each reactant.

| Id   | Name | SBO |
|------|------|-----|
| pSOS |      |     |

#### **Product**

Table 36: Properties of each product.

| Id  | Name | SBO |
|-----|------|-----|
| SOS |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{11} = \text{vol}(c1) \cdot \text{J}10_{-}\text{k} \cdot [pSOS]$$
 (22)

Table 37: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J10_k |      | 0.002          |          |

# 5.12 Reaction J11

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

Name pSOS\_Grb2\_dephosphorylation

# **Reaction equation**

$$pSOS\_Grb2 \longrightarrow SOS\_Grb2$$
 (23)

# Reactant

Table 38: Properties of each reactant.

| Id        | Name | SBO |
|-----------|------|-----|
| pSOS_Grb2 |      |     |

#### **Product**

Table 39: Properties of each product.

| Id       | Name | SBO |
|----------|------|-----|
| SOS_Grb2 |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{12} = \text{vol}(c1) \cdot \text{J11\_k} \cdot [pSOS\_Grb2]$$
 (24)

Table 40: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J11_k |      | 0.002          |          |

#### 5.13 Reaction J12

This is a reversible reaction of two reactants forming one product.

Name binding\_Shc\_LdpEGFR

# **Reaction equation**

$$L_dpEGFR + Shc \Longrightarrow Shc_dpEGFR$$
 (25)

#### **Reactants**

Table 41: Properties of each reactant.

| Id              | Name | SBO |
|-----------------|------|-----|
| L_dpEGFR<br>Shc |      |     |

#### **Product**

Table 42: Properties of each product.

| Id         | Name | SBO |
|------------|------|-----|
| Shc_dpEGFR |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{13} = \text{vol}(c1) \cdot (\text{J}12\_\text{k}1 \cdot [\text{L\_dpEGFR}] \cdot [\text{Shc}] - \text{J}12\_\text{k}2 \cdot [\text{Shc\_dpEGFR}])$$
 (26)

Table 43: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant  |
|--------|------|----------------|-----------|
| J12_k1 |      | 10.0           |           |
| J12_k2 |      | 0.2            | $\square$ |

# **5.14 Reaction** J13

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

Name Shc\_dpEGFR\_phosphorylation

# **Reaction equation**

$$Shc\_dpEGFR \longrightarrow pShc\_dpEGFR$$
 (27)

#### Reactant

Table 44: Properties of each reactant.

| Id         | Name | SBO |
|------------|------|-----|
| Shc_dpEGFR |      |     |

Table 45: Properties of each product.

| Id          | Name | SBO |
|-------------|------|-----|
| pShc_dpEGFR |      |     |

**Derived unit** contains undeclared units

$$v_{14} = \text{vol}(c1) \cdot \text{J}13\_k \cdot [\text{Shc\_dpEGFR}]$$
 (28)

Table 46: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J13_k |      | 1.0            |          |

#### 5.15 Reaction J14

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

Name dpEGFR\_c\_Cbl\_ubiquitination

# **Reaction equation**

$$dpEGFR\_c\_Cbl \longrightarrow dpEGFR\_c\_Cbl\_ubiq$$
 (29)

#### Reactant

Table 47: Properties of each reactant.

| Id           | Name | SBO |
|--------------|------|-----|
| dpEGFR_c_Cbl |      |     |

Table 48: Properties of each product.

| Id                | Name | SBO |
|-------------------|------|-----|
| dpEGFR_c_Cbl_ubiq |      |     |

**Derived unit** contains undeclared units

$$v_{15} = \text{vol}(c1) \cdot \text{J}14\_k \cdot [\text{dpEGFR\_c\_Cbl}]$$
(30)

Table 49: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J14_k |      | 0.05           |          |

#### 5.16 Reaction J15

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

Name dpEGFR\_cCbl\_degrad

# **Reaction equation**

$$dpEGFR\_c\_Cbl\_ubiq \longrightarrow proteosome + c\_Cbl$$
 (31)

#### Reactant

Table 50: Properties of each reactant.

| Id                | Name | SBO |
|-------------------|------|-----|
| dpEGFR_c_Cbl_ubiq |      |     |

## **Products**

Table 51: Properties of each product.

| Id                  | Name       | SBO |
|---------------------|------------|-----|
| proteosome<br>c_Cbl | proteasome |     |

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{16} = \text{vol}(c1) \cdot \text{J}15\_k \cdot [\text{dpEGFR\_c\_Cbl\_ubiq}]$$
(32)

Table 52: Properties of each parameter.

| Id    | Name | SBO | Value | Unit | Constant |
|-------|------|-----|-------|------|----------|
| J15_k |      |     | 0.001 |      |          |

# 5.17 Reaction J16

This is a reversible reaction of two reactants forming one product.

Name binding\_cCbl\_Shc\_dpEGFR

# **Reaction equation**

$$c_Cbl + Shc_dpEGFR \Longrightarrow Shc_dpEGFR_cCbl$$
 (33)

#### **Reactants**

Table 53: Properties of each reactant.

| Id         | Name | SBO |
|------------|------|-----|
| c_Cbl      |      |     |
| Shc_dpEGFR |      |     |

#### **Product**

Table 54: Properties of each product.

| Id               | Name | SBO |
|------------------|------|-----|
| Shc_dpEGFR_c_Cbl |      |     |

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{17} = \text{vol}(c1) \cdot (\text{J}16\_\text{k}1 \cdot [c\_\text{Cbl}] \cdot [\text{Shc\_dpEGFR}] - \text{J}16\_\text{k}2 \cdot [\text{Shc\_dpEGFR\_c\_Cbl}]) \quad (34)$$

Table 55: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant       |
|--------|------|----------------|----------------|
| J16_k1 |      | 0.5            | $\blacksquare$ |
| J16_k2 |      | 0.2            | $\square$      |

#### 5.18 Reaction J17

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

Name Shc\_dpEGFR\_c\_CBl\_Ubiquitination

# **Reaction equation**

$$Shc\_dpEGFR\_c\_Cbl \longrightarrow Shc\_dpEGFR\_c\_Cbl\_ubiq$$
 (35)

#### Reactant

Table 56: Properties of each reactant.

Id Name SBO

Shc\_dpEGFR\_c\_Cbl

#### **Product**

Table 57: Properties of each product.

| Id                    | Name | SBO |
|-----------------------|------|-----|
| Shc_dpEGFR_c_Cbl_ubiq |      |     |

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{18} = \text{vol}(c1) \cdot \text{J}17_{\text{-k}} \cdot [\text{Shc\_dpEGFR\_c\_Cbl}]$$
(36)

Table 58: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J17_k |      | 0.05           |          |

#### 5.19 Reaction J18

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

Name Shc\_dpEGFR\_c\_Cbl\_ubiq\_Degradation

# **Reaction equation**

$$Shc\_dpEGFR\_c\_Cbl\_ubiq \longrightarrow proteosome + c\_Cbl + Shc$$
 (37)

#### Reactant

Table 59: Properties of each reactant.

| ruble 33. I roperties of each reactant. |      |     |  |  |
|---|------|-----|--|--|
| Id                                      | Name | SBO |  |  |
| Shc_dpEGFR_c_Cbl_ubiq                   |      |     |  |  |

#### **Products**

Table 60: Properties of each product.

| Id                         | Name       | SBO |
|----------------------------|------------|-----|
| proteosome<br>c_Cbl<br>Shc | proteasome |     |

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{19} = \text{vol}(c1) \cdot \text{J}18\_\text{k} \cdot [\text{Shc\_dpEGFR\_c\_Cbl\_ubiq}]$$
 (38)

Table 61: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J18_k |      | 0.001          | Ø        |

#### 5.20 Reaction J19

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

 $\textbf{Name} \hspace{0.2cm} pShc\_dpEGFR\_c\_Cbl\_ubiquitination$ 

# **Reaction equation**

$$pShc\_dpEGFR\_c\_Cbl \longrightarrow pShc\_dpEGFR\_c\_Cbl\_ubiq$$
 (39)

#### Reactant

Table 62: Properties of each reactant.

| Tuble 02: Troperties 0 | i cacii ici | actuiit. |
|------------------------|-------------|----------|
| Id                     | Name        | SBO      |
| pShc_dpEGFR_c_Cbl      |             |          |

#### **Product**

Table 63: Properties of each product.

| Id                     | Name | SBO |
|------------------------|------|-----|
| pShc_dpEGFR_c_Cbl_ubiq |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{20} = \text{vol}(c1) \cdot \text{J}19\_\text{k} \cdot [\text{pShc\_dpEGFR\_c\_Cbl}]$$
 (40)

Table 64: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J19_k |      | 0.05           |          |

# **5.21 Reaction** J20

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

Name pShc\_dpEGFR\_c\_Cbl\_ubiq\_degradation

# **Reaction equation**

$$pShc\_dpEGFR\_c\_Cbl\_ubiq \longrightarrow proteosome + c\_Cbl + pShc$$
 (41)

#### Reactant

Table 65: Properties of each reactant.

| Id                     | Name | SBO |
|------------------------|------|-----|
| pShc_dpEGFR_c_Cbl_ubiq |      |     |

#### **Products**

Table 66: Properties of each product.

| 1                           | 1          |     |
|-----------------------------|------------|-----|
| Id                          | Name       | SBO |
| proteosome<br>c_Cbl<br>pShc | proteasome |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{21} = \text{vol}(c1) \cdot \text{J20\_k} \cdot [\text{pShc\_dpEGFR\_c\_Cbl\_ubiq}]$$
 (42)

Table 67: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J20_k |      | 0.001          |          |

#### **5.22 Reaction** J21

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

Name Shc\_dpEGFR\_c\_Cblphosphorylation

# **Reaction equation**

$$Shc\_dpEGFR\_c\_Cbl \longrightarrow pShc\_dpEGFR\_c\_Cbl$$
 (43)

#### Reactant

Table 68: Properties of each reactant.

| Id               | Name | SBO |
|------------------|------|-----|
| Shc_dpEGFR_c_Cbl |      |     |

| Table 07. Troperties 0 | r cacii pi | oduct. |
|------------------------|------------|--------|
| Id                     | Name       | SBO    |
| pShc_dpEGFR_c_Cbl      |            |        |

Derived unit contains undeclared units

$$v_{22} = \text{vol}(c1) \cdot \text{J21\_k} \cdot [\text{Shc\_dpEGFR\_c\_Cbl}]$$
(44)

Table 70: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J21_k |      | 1.0            |          |

#### **5.23 Reaction** J22

This is a reversible reaction of two reactants forming one product.

Name binding\_Grb2\_SOS\_pShc

# **Reaction equation**

$$pShc + SOS\_Grb2 \Longrightarrow Grb2\_SOS\_pShc$$
 (45)

#### **Reactants**

Table 71: Properties of each reactant.

| Id               | Name | SBO |
|------------------|------|-----|
| pShc<br>SOS_Grb2 |      |     |

Table 72: Properties of each product.

| Id            | Name | SBO |
|---------------|------|-----|
| Grb2_SOS_pShc |      |     |

Derived unit contains undeclared units

$$v_{23} = \text{vol}(c1) \cdot (\text{J22\_k1} \cdot [\text{pShc}] \cdot [\text{SOS\_Grb2}] - \text{J22\_k2} \cdot [\text{Grb2\_SOS\_pShc}])$$
 (46)

Table 73: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J22_k1 |      | 10.0           |          |
| J22_k2 |      | 0.2            |          |

#### 5.24 Reaction J23

This is a reversible reaction of two reactants forming one product.

Name binding\_Grb2\_SOS\_pShc\_dpEGFR

# **Reaction equation**

$$L\_dpEGFR + Grb2\_SOS\_pShc \Longrightarrow Grb2\_SOS\_pShc\_dpEGFR$$
 (47)

## **Reactants**

Table 74: Properties of each reactant.

| Id            | Name | SBO |
|---------------|------|-----|
| L_dpEGFR      |      |     |
| Grb2_SOS_pShc |      |     |

#### **Product**

Table 75: Properties of each product.

| Id                   | Name | SBO |
|----------------------|------|-----|
| Grb2_SOS_pShc_dpEGFR |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{24} = \text{vol} (\text{c1}) \cdot (\text{J23\_k1} \cdot [\text{L\_dpEGFR}] \cdot [\text{Grb2\_SOS\_pShc}] - \text{J23\_k2} \cdot [\text{Grb2\_SOS\_pShc\_dpEGFR}])$$

$$(48)$$

Table 76: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J23_k1 |      | 10.0           |          |
| J23_k2 |      | 0.2            |          |

### 5.25 Reaction J24

This is a reversible reaction of two reactants forming one product.

 $\textbf{Name} \ binding\_Grb2\_SOS\_pShc\_dpEGFR$ 

## **Reaction equation**

$$pShc\_dpEGFR + SOS\_Grb2 \Longrightarrow Grb2\_SOS\_pShc\_dpEGFR$$
 (49)

#### **Reactants**

Table 77: Properties of each reactant.

| Id                      | Name | SBO |
|-------------------------|------|-----|
| pShc_dpEGFR<br>SOS_Grb2 |      |     |

### **Product**

Table 78: Properties of each product.

| Id                   | Name | SBO |
|----------------------|------|-----|
| Grb2_SOS_pShc_dpEGFR |      |     |

#### **Kinetic Law**

$$v_{25} = \text{vol}\,(\text{c1}) \cdot (\text{J24\_k1} \cdot [\text{pShc\_dpEGFR}] \cdot [\text{SOS\_Grb2}] - \text{J24\_k2} \cdot [\text{Grb2\_SOS\_pShc\_dpEGFR}])$$
(50)

Table 79: Properties of each parameter.

| Id       | Name | SBO Value Unit | Constant  |
|----------|------|----------------|-----------|
| J24_k1   |      | 10.0           | $\square$ |
| $J24_k2$ |      | 0.2            |           |

### 5.26 Reaction J25

This is a reversible reaction of two reactants forming one product.

Name binding\_c\_Cbl\_Grb2\_SOS\_pShc\_dpEGFR

## **Reaction equation**

$$c\_Cbl + Grb2\_SOS\_pShc\_dpEGFR \Longrightarrow Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl$$
 (51)

#### **Reactants**

Table 80: Properties of each reactant.

| Id                            | Name | SBO |
|-------------------------------|------|-----|
| c_Cbl<br>Grb2_SOS_pShc_dpEGFR |      |     |

#### **Product**

Table 81: Properties of each product.

| Id                         | Name | SBO |
|----------------------------|------|-----|
| Grb2_SOS_pShc_dpEGFR_c_Cbl |      |     |

#### **Kinetic Law**

$$v_{26} = \text{vol}(c1) \cdot (\text{J25\_k1} \cdot [\text{c\_Cbl}] \cdot [\text{Grb2\_SOS\_pShc\_dpEGFR}] - \text{J25\_k2}$$

$$\cdot [\text{Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl}])$$
(52)

Table 82: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J25_k1 |      | 0.5            |          |
| J25_k2 |      | 0.2            |          |

### **5.27 Reaction** J27

This is a reversible reaction of two reactants forming one product.

Name binding\_Grb2\_SOS\_pShc\_to\_dpEGFR\_c\_Cbl

# **Reaction equation**

$$dpEGFR\_c\_Cbl + Grb2\_SOS\_pShc \Longrightarrow Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl$$
 (53)

#### **Reactants**

Table 83: Properties of each reactant.

| Id            | Name | SBO |
|---------------|------|-----|
| dpEGFR_c_Cbl  |      |     |
| Grb2_SOS_pShc |      |     |

#### **Product**

Table 84: Properties of each product.

| Id                         | Name | SBO |
|----------------------------|------|-----|
| Grb2_SOS_pShc_dpEGFR_c_Cbl |      |     |

#### **Kinetic Law**

$$v_{27} = \text{vol}(c1) \cdot (\text{J27\_k1} \cdot [\text{dpEGFR\_c\_Cbl}] \cdot [\text{Grb2\_SOS\_pShc}] - \text{J27\_k2}$$

$$\cdot [\text{Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl}])$$
(54)

Table 85: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant   |
|--------|------|----------------|------------|
| J27_k1 |      | 10.0           | lacksquare |
| J27_k2 |      | 0.2            |            |

### 5.28 Reaction J28

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

Name Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl\_ubiquitination

# **Reaction equation**

 $Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl \longrightarrow Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl\_ubiq \qquad (55)$ 

#### Reactant

Table 86: Properties of each reactant.

| Id                         | Name |  |
|----------------------------|------|--|
| Grb2_SOS_pShc_dpEGFR_c_Cbl |      |  |

#### **Product**

Table 87: Properties of each product.

| Id                              | Name | SBO |
|---------------------------------|------|-----|
| Grb2_SOS_pShc_dpEGFR_c_Cbl_ubiq |      |     |

#### **Kinetic Law**

$$v_{28} = \text{vol}(c1) \cdot \text{J28\_k} \cdot [\text{Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl}]$$
 (56)

Table 88: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant     |
|-------|------|----------------|--------------|
| J28_k |      | 0.05           | $\checkmark$ |

### 5.29 Reaction J29

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

Name Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl\_ubiq\_degradation

## **Reaction equation**

$$Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl\_ubiq \longrightarrow proteosome + c\_Cbl + Grb2\_SOS\_pShc \qquad (57)$$

### Reactant

Table 89: Properties of each reactant.

| Id                              | Name | SBO |
|---------------------------------|------|-----|
| Grb2_SOS_pShc_dpEGFR_c_Cbl_ubiq |      |     |

#### **Products**

Table 90: Properties of each product.

| Id                                   | Name       | SBO |
|--------------------------------------|------------|-----|
| proteosome<br>c_Cbl<br>Grb2_SOS_pShc | proteasome |     |

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{29} = \text{vol}(c1) \cdot \text{J29\_k} \cdot [\text{Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl\_ubiq}]$$
 (58)

Table 91: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J29_k |      | 0.001          | Ø        |

#### 5.30 Reaction J30

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

Name Grb2\_SOS\_pShc\_Dissociation

## **Reaction equation**

$$Grb2\_SOS\_pShc \longrightarrow Shc + SOS\_Grb2$$
 (59)

### Reactant

Table 92: Properties of each reactant.

| Id            | Name | SBO |
|---------------|------|-----|
| Grb2_SOS_pShc |      |     |

### **Products**

Table 93: Properties of each product.

| Id          | Name | SBO |
|-------------|------|-----|
| Shc         |      |     |
| $SOS\_Grb2$ |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{30} = J30_{k} \cdot [Grb2\_SOS\_pShc]$$
 (60)

Table 94: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant  |
|-------|------|----------------|-----------|
| J30_k |      | 0.005          | $\square$ |

#### 5.31 Reaction J31

This is an irreversible reaction of one reactant forming one product influenced by 13 modifiers.

# **Reaction equation**

Dok L\_dpEGFR, Shc\_dpEGFR, pShc\_dpEGFR, Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl, Grb2\_SOS\_pShc\_dpEGFR, dpEGFR, dpEGF

Table 95: Properties of each reactant.

| Id  | Name | SBO |
|-----|------|-----|
| Dok |      |     |

## **Modifiers**

Table 96: Properties of each modifier.

| Id                         | Name | SBO |
|----------------------------|------|-----|
| L_dpEGFR                   |      |     |
| ${	t Shc\_dpEGFR}$         |      |     |
| $pShc_dpEGFR$              |      |     |
| Grb2_SOS_pShc_dpEGFR_c_Cbl |      |     |
| Grb2_SOS_pShc_dpEGFR       |      |     |
| dpEGFR_c_Cbl               |      |     |
| Shc_dpEGFR_c_Cbl           |      |     |
| pShc_dpEGFR_c_Cbl          |      |     |
| FRS2_dpEGFR                |      |     |
| pFRS2_dpEGFR               |      |     |
| Crk_C3G_pFRS2_dpEGFR       |      |     |
| FRS2_dpEGFR_c_Cbl          |      |     |
| Crk_C3G_pFRS2_dpEGFR_c_Cbl |      |     |

## **Product**

Table 97: Properties of each product.

| Id   | Name | SBO |
|------|------|-----|
| pDok |      |     |

## **Kinetic Law**

$$v_{31} = vol(c1)$$
 
$$J31\_Vmax \cdot [Dok] \cdot ([L\_dpEGFR] + [Shc\_dpEGFR] + [pShc\_dpEGFR] + [Grb2\_SOS\_pShc\_dpEGFR] + [dpEGFR] +$$

Table 98: Properties of each parameter.

| Id             | Name | SBO Value Unit | Constant |
|----------------|------|----------------|----------|
| J31_Vmax       |      | 0.2            |          |
| $\rm J31\_Km1$ |      | 0.1            |          |

## **5.32 Reaction** J32

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

Name pShc\_dephosphorylation

# **Reaction equation**

$$pShc \longrightarrow Shc \tag{63}$$

#### Reactant

Table 99: Properties of each reactant.

| Id   | Name | SBO |
|------|------|-----|
| pShc |      |     |

# **Product**

Table 100: Properties of each product.

| Id  | Name | SBO |
|-----|------|-----|
| Shc |      |     |

#### **Kinetic Law**

$$v_{32} = \text{vol}(c1) \cdot \text{J32\_k} \cdot [\text{pShc}] \tag{64}$$

Table 101: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J32_k |      | 0.005          |          |

## 5.33 Reaction J33

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

Name pFRS2\_dephosphorylation

## **Reaction equation**

$$pFRS2 \longrightarrow FRS2 \tag{65}$$

### Reactant

Table 102: Properties of each reactant.

| Id    | Name | SBO |
|-------|------|-----|
| pFRS2 |      |     |

### **Product**

Table 103: Properties of each product.

| Id   | Name | SBO |
|------|------|-----|
| FRS2 |      |     |

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{33} = \text{vol}(c1) \cdot \text{J33\_k} \cdot [\text{pFRS2}] \tag{66}$$

Table 104: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J33_k |      | 0.005          |          |

## 5.34 Reaction J34

This is a reversible reaction of two reactants forming one product.

Name binding\_Crk\_to\_C3G

## **Reaction equation**

$$Crk + C3G \rightleftharpoons Crk\_C3G$$
 (67)

#### **Reactants**

Table 105: Properties of each reactant.

| Id  | Name | SBO |
|-----|------|-----|
| Crk |      |     |
| C3G |      |     |

#### **Product**

Table 106: Properties of each product.

| Id      | Name | SBO |
|---------|------|-----|
| Crk_C3G |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{34} = \text{vol}(c1) \cdot (J34\_k1 \cdot [Crk] \cdot [C3G] - J34\_k2 \cdot [Crk\_C3G])$$
 (68)

Table 107: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant     |
|--------|------|----------------|--------------|
| J34_k1 |      | 1.000          |              |
| J34_k2 |      | 0.002          | $\checkmark$ |

### 5.35 Reaction J35

This is a reversible reaction of two reactants forming one product.

Name binding\_L\_dpEGFR\_to\_FRS2

## **Reaction equation**

$$L\_dpEGFR + FRS2 \Longrightarrow FRS2\_dpEGFR$$
 (69)

#### **Reactants**

Table 108: Properties of each reactant.

| Id               | Name | SBO |
|------------------|------|-----|
| L_dpEGFR<br>FRS2 |      |     |

#### **Product**

Table 109: Properties of each product.

| Id          | Name | SBO |
|-------------|------|-----|
| FRS2_dpEGFR |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{35} = \text{vol}(c1) \cdot (\text{J35\_k1} \cdot [\text{L\_dpEGFR}] \cdot [\text{FRS2}] - \text{J35\_k2} \cdot [\text{FRS2\_dpEGFR}])$$
 (70)

Table 110: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant  |
|--------|------|----------------|-----------|
| J35_k1 |      | 1.0            | Ø         |
| J35_k2 |      | 0.2            | $\square$ |

### 5.36 Reaction J36

This is a reversible reaction of two reactants forming one product.

Name binding\_pFRS2\_to\_L\_dpEGFR

## **Reaction equation**

$$L\_dpEGFR + pFRS2 \Longrightarrow pFRS2\_dpEGFR \tag{71}$$

Table 111: Properties of each reactant.

| Id       | Name | SBO |
|----------|------|-----|
| L_dpEGFR |      |     |
| pFRS2    |      |     |

Table 112: Properties of each product.

| Id           | Name | SBO |
|--------------|------|-----|
| pFRS2_dpEGFR |      |     |

#### **Kinetic Law**

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

$$v_{36} = \text{vol}(c1) \cdot (\text{J36\_k1} \cdot [\text{L\_dpEGFR}] \cdot [\text{pFRS2}] - \text{J36\_k2} \cdot [\text{pFRS2\_dpEGFR}])$$
 (72)

Table 113: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J36_k1 |      | 1.0            | $lue{2}$ |
| J36_k2 |      | 0.2            |          |

# 5.37 Reaction J37

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

 $\textbf{Name} \ \ FRS2\_dpEGFR phsphorylation$ 

## **Reaction equation**

$$FRS2\_dpEGFR \longrightarrow pFRS2\_dpEGFR \tag{73}$$

Table 114: Properties of each reactant.

| Id          | Name | SBO |
|-------------|------|-----|
| FRS2_dpEGFR |      |     |

Table 115: Properties of each product.

| Id           | Name | SBO |
|--------------|------|-----|
| pFRS2_dpEGFR |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{37} = \text{vol}(c1) \cdot \text{J37\_k} \cdot [\text{FRS2\_dpEGFR}]$$
 (74)

Table 116: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J37_k |      | 1.0            |          |

### 5.38 Reaction J38

This is a reversible reaction of two reactants forming one product.

Name binding\_Crk\_C3G\_to\_pFRS2\_pRTK

## **Reaction equation**

$$pFRS2\_dpEGFR + Crk\_C3G \Longrightarrow Crk\_C3G\_pFRS2\_dpEGFR$$
 (75)

## Reactants

Table 117: Properties of each reactant.

| Id                      | Name | SBO |
|-------------------------|------|-----|
| pFRS2_dpEGFR<br>Crk_C3G |      |     |

Table 118: Properties of each product.

| Id                   | Name |  |
|----------------------|------|--|
| Crk_C3G_pFRS2_dpEGFR |      |  |

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

$$v_{38} = vol(c1) \cdot (J38\_k1 \cdot [pFRS2\_dpEGFR] \cdot [Crk\_C3G] - J38\_k2 \cdot [Crk\_C3G\_pFRS2\_dpEGFR])$$
(76)

Table 119: Properties of each parameter.

| Id     | Name | SBO Value | Unit | Constant |
|--------|------|-----------|------|----------|
| J38_k1 |      | 1.0       |      |          |
| J38_k2 |      | 0.2       |      |          |

### 5.39 Reaction J39

This is a reversible reaction of two reactants forming one product.

Name binding\_c\_Cbl\_to\_FRS2\_dpEGFR

# **Reaction equation**

$$FRS2\_dpEGFR + c\_Cbl \Longrightarrow FRS2\_dpEGFR\_c\_Cbl$$
 (77)

#### **Reactants**

Table 120: Properties of each reactant.

| Id          | Name | SBO |
|-------------|------|-----|
| FRS2_dpEGFR |      |     |
| c_Cbl       |      |     |

Table 121: Properties of each product.

| Table 121. I Toperties of each product. |      |     |  |
|---|------|-----|--|
| Id                                      | Name | SBO |  |
| FRS2_dpEGFR_c_Cbl                       |      |     |  |

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

$$v_{39} = vol(c1) \cdot (J39\_k1 \cdot [FRS2\_dpEGFR] \cdot [c\_Cbl] - J39\_k2 \cdot [FRS2\_dpEGFR\_c\_Cbl]) \quad (78)$$

Table 122: Properties of each parameter.

|        |      | 1 1            |           |
|--------|------|----------------|-----------|
| Id     | Name | SBO Value Unit | Constant  |
| J39_k1 |      | 0.5            |           |
| J39_k2 |      | 0.2            | $\square$ |

### 5.40 Reaction J40

This is a reversible reaction of two reactants forming one product.

Name binding\_c\_Cbl\_to\_pFRS2\_dpEGFR

## **Reaction equation**

$$c\_Cbl + pFRS2\_dpEGFR \Longrightarrow pFRS2\_dpEGFR\_c\_Cbl$$
 (79)

#### **Reactants**

Table 123: Properties of each reactant.

| Id           | Name | SBO |
|--------------|------|-----|
| c_Cbl        |      |     |
| pFRS2_dpEGFR |      |     |

Table 124: Properties of each product.

| Id                 | Name | SBO |
|--------------------|------|-----|
| pFRS2_dpEGFR_c_Cbl |      |     |

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

$$v_{40} = \text{vol}\left(\text{c1}\right) \cdot \left(\text{J40\_k1} \cdot \left[\text{c\_Cbl}\right] \cdot \left[\text{pFRS2\_dpEGFR}\right] - \text{J40\_k2} \cdot \left[\text{pFRS2\_dpEGFR\_c\_Cbl}\right]\right) \quad (80)$$

Table 125: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J40_k1 |      | 0.5            |          |
| J40_k2 |      | 0.2            |          |

### **5.41 Reaction** J41

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

Name pFRS2\_dpEGFR\_c\_Cbl\_ubiquitiation

## **Reaction equation**

$$pFRS2\_dpEGFR\_c\_Cbl \longrightarrow pFRS2\_dpEGFR\_c\_Cbl\_ubiq$$
 (81)

## Reactant

Table 126: Properties of each reactant.

| Id                 | Name | SBO |
|--------------------|------|-----|
| pFRS2_dpEGFR_c_Cbl |      |     |

## **Product**

Table 127: Properties of each product.

| Id                      | Name | SBO |
|-------------------------|------|-----|
| pFRS2_dpEGFR_c_Cbl_ubiq |      |     |

## **Kinetic Law**

$$v_{41} = \text{vol}(c1) \cdot \text{J41\_k} \cdot [\text{pFRS2\_dpEGFR\_c\_Cbl}]$$
 (82)

Table 128: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J41_k |      | 0.05           |          |

### 5.42 Reaction J42

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

Name FRS2\_dpEGFR\_c\_Cbl\_ubiquitination

# **Reaction equation**

$$FRS2\_dpEGFR\_c\_Cbl \longrightarrow FRS2\_dpEGFR\_c\_Cbl\_ubiq$$
 (83)

#### Reactant

Table 129: Properties of each reactant.

| Id                | Name | SBO |
|-------------------|------|-----|
| FRS2_dpEGFR_c_Cbl |      |     |

## **Product**

Table 130: Properties of each product.

| Id                     | Name | SBO |
|------------------------|------|-----|
| FRS2_dpEGFR_c_Cbl_ubiq |      |     |

#### **Kinetic Law**

$$v_{42} = \text{vol}(c1) \cdot \text{J42\_k} \cdot [\text{FRS2\_dpEGFR\_c\_Cbl}]$$
 (84)

Table 131: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J42_k |      | 0.05           |          |

### 5.43 Reaction J43

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

Name FRS2\_dpEGFR\_c\_Cbl\_phosphorylation

## **Reaction equation**

$$FRS2\_dpEGFR\_c\_Cbl \longrightarrow pFRS2\_dpEGFR\_c\_Cbl$$
 (85)

### Reactant

Table 132: Properties of each reactant.

Id Name SBO

FRS2\_dpEGFR\_c\_Cbl

### **Product**

Table 133: Properties of each product.

Id Name SBO

pFRS2\_dpEGFR\_c\_Cbl

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{43} = \text{vol}(c1) \cdot \text{J43\_k} \cdot [\text{FRS2\_dpEGFR\_c\_Cbl}]$$
 (86)

Table 134: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J43_k |      | 1.0            |          |

### 5.44 Reaction J44

This is a reversible reaction of two reactants forming one product.

Name binding\_Crk\_C3G\_to\_pFRS2\_pFRS2\_dpEGFR\_c\_Cbl

## **Reaction equation**

$$pFRS2\_dpEGFR\_c\_Cbl + Crk\_C3G \Longrightarrow Crk\_C3G\_pFRS2\_dpEGFR\_c\_Cbl$$
 (87)

#### **Reactants**

Table 135: Properties of each reactant.

| Id                            | Name | SBO |
|-------------------------------|------|-----|
| pFRS2_dpEGFR_c_Cbl<br>Crk_C3G |      |     |

#### **Product**

Table 136: Properties of each product.

| Id                     | Name | SBO |
|------------------------|------|-----|
| Crk_C3G_pFRS2_dpEGFR_c | _Cbl |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{44} = \text{vol}(c1) \cdot (\text{J}44\_\text{k}1 \cdot [\text{pFRS2\_dpEGFR\_c\_Cbl}] \cdot [\text{Crk\_C3G}] - \text{J}44\_\text{k}2$$

$$\cdot [\text{Crk\_C3G\_pFRS2\_dpEGFR\_c\_Cbl}])$$
(88)

Table 137: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant       |
|--------|------|----------------|----------------|
| J44_k1 |      | 1.0            | $\overline{Z}$ |
| J44_k2 |      | 0.2            |                |

### 5.45 Reaction J45

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

Name Crk\_C3G\_pFRS2\_dpEGFR\_c\_Cbl\_ubiquitination

## **Reaction equation**

$$Crk_C3G_pFRS2\_dpEGFR\_c\_Cbl\_ \longrightarrow Crk_C3G_pFRS2\_dpEGFR\_c\_Cbl\_ubiq$$
 (89)

#### Reactant

Table 138: Properties of each reactant

|     | Tuble 130. I repetites of each reactant. |      |     |  |  |
|-----|--|------|-----|--|--|
| Id  |  | Name | SBO |  |  |
| Crk | _C3G_pFRS2_dpEGFR_c_Cbl                  |      |     |  |  |

### **Product**

Table 139: Properties of each product.

| Id                 |              |   | Name | SBO |
|--------------------|--------------|---|------|-----|
| Crk_C3G_pFRS2_dpEG | FR_c_Cbl_ubi | q |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{45} = \text{vol}(c1) \cdot \text{J45\_k} \cdot [\text{Crk\_C3G\_pFRS2\_dpEGFR\_c\_Cbl}]$$

$$\tag{90}$$

Table 140: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J45_k |      | 0.05           |          |

## **5.46 Reaction** J46

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

Name FRS2\_dpEGFR\_c\_Cbl\_ubiq\_dissociation

## **Reaction equation**

$$FRS2\_dpEGFR\_c\_Cbl\_ubiq \longrightarrow proteosome + c\_Cbl + FRS2$$
 (91)

Table 141: Properties of each reactant.

| Id                     | Name | SBO |
|------------------------|------|-----|
| FRS2_dpEGFR_c_Cbl_ubiq |      |     |

Table 142: Properties of each product.

| Id                          | Name       | SBO |
|-----------------------------|------------|-----|
| proteosome<br>c_Cbl<br>FRS2 | proteasome |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{46} = \text{vol}(c1) \cdot \text{J46\_k} \cdot [\text{FRS2\_dpEGFR\_c\_Cbl\_ubiq}]$$
 (92)

Table 143: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J46_k |      | 0.001          |          |

# **5.47 Reaction** J47

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

Name pFRS2\_dpEGFR\_c\_Cbl\_ubiq\_dissociation

## **Reaction equation**

$$pFRS2\_dpEGFR\_c\_Cbl\_ubiq \longrightarrow proteosome + c\_Cbl + pFRS2$$
 (93)

### Reactant

Table 144: Properties of each reactant.

| Tuble 111. 11operties of each reactant. |      |     |  |  |
|---|------|-----|--|--|
| Id                                      | Name | SBO |  |  |
| pFRS2_dpEGFR_c_Cbl_ubiq                 |      |     |  |  |

Table 145: Properties of each product.

| Id                           | Name       | SBO |
|------------------------------|------------|-----|
| proteosome<br>c_Cbl<br>pFRS2 | proteasome |     |

**Derived unit** contains undeclared units

$$v_{47} = \text{vol}(c1) \cdot \text{J47\_k} \cdot [\text{pFRS2\_dpEGFR\_c\_Cbl\_ubiq}]$$
 (94)

Table 146: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J47_k |      | 0.001          |          |

## 5.48 Reaction J49

This is a reversible reaction of two reactants forming one product.

Name binding\_RasGAP\_to\_pDOK

# **Reaction equation**

$$pDok + RasGAP \Longrightarrow pDok\_RasGAP$$
 (95)

### **Reactants**

Table 147: Properties of each reactant.

| Id             | Name | SBO |
|----------------|------|-----|
| pDok<br>RasGAP |      |     |

Table 148: Properties of each product.

| Id          | Name | SBO |
|-------------|------|-----|
| pDok_RasGAP |      |     |

**Derived unit** contains undeclared units

$$v_{48} = vol\left(c1\right) \cdot \left(J49\_k1 \cdot [pDok] \cdot [RasGAP] - J49\_k2 \cdot [pDok\_RasGAP]\right) \tag{96}$$

Table 149: Properties of each parameter.

|        |      | 1 1            |                |
|--------|------|----------------|----------------|
| Id     | Name | SBO Value Unit | Constant       |
| J49_k1 |      | 0.12           | $\overline{Z}$ |
| J49_k2 |      | 0.01           | $\square$      |

### 5.49 Reaction J50

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

Name SOS\_Grb2\_phosphorylation

# **Reaction equation**

$$SOS\_Grb2 \xrightarrow{dppERK} pSOS\_Grb2$$
 (97)

### Reactant

Table 150: Properties of each reactant.

| Id       | Name | SBO |
|----------|------|-----|
| SOS_Grb2 |      |     |

#### **Modifier**

Table 151: Properties of each modifier.

| Id     | Name | SBO |
|--------|------|-----|
| dppERK |      |     |

Table 152: Properties of each product.

| Id        | Name | SBO |
|-----------|------|-----|
| pSOS_Grb2 |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{49} = \text{vol}(c1) \cdot \frac{\text{J50\_Vmax} \cdot [\text{SOS\_Grb2}] \cdot [\text{dppERK}]}{\text{J50\_Km1} + [\text{SOS\_Grb2}]}$$
(98)

Table 153: Properties of each parameter.

| Id               | Name | SBO Value Unit | Constant       |
|------------------|------|----------------|----------------|
| J50_Vmax         |      | 1.000          | $ \mathbf{Z} $ |
| ${\tt J50\_Km1}$ |      | 25.641         | $\checkmark$   |

## 5.50 Reaction J51

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

Name SOS\_phosphorylation

### **Reaction equation**

$$SOS \xrightarrow{dppERK} pSOS$$
 (99)

#### Reactant

Table 154: Properties of each reactant.

| Id  | Name | SBO |
|-----|------|-----|
| SOS |      |     |

## **Modifier**

Table 155: Properties of each modifier.

| Id     | Name | SBO |
|--------|------|-----|
| dppERK |      |     |

Table 156: Properties of each product.

| Id   | Name | SBO |
|------|------|-----|
| pSOS |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{50} = \text{vol}(c1) \cdot \frac{\text{J51\_Vmax} \cdot [\text{SOS}] \cdot [\text{dppERK}]}{\text{J51\_Km1} + [\text{SOS}]}$$
(100)

Table 157: Properties of each parameter.

| Id             | Name | SBO V | alue Unit | Constant |
|----------------|------|-------|-----------|----------|
| $J51_{Vmax}$   |      | 1     | 1.000     |          |
| $\tt J51\_Km1$ |      | 25    | 5.641     |          |

### 5.51 Reaction J52

This is a reversible reaction of two reactants forming one product.

Name binding\_c\_Raf\_to\_Ras\_GTP

# **Reaction equation**

$$c_Raf + Ras_GTP \Longrightarrow c_Raf_Ras_GTP$$
 (101)

Table 158: Properties of each reactant.

| Id    | Name | SBO |
|-------|------|-----|
| c_Raf |      |     |

| Id      | Name | SBO |
|---------|------|-----|
| Ras_GTP |      |     |

Table 159: Properties of each product.

| Id            | Name | SBO |
|---------------|------|-----|
| c_Raf_Ras_GTP |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{51} = \text{vol}(c1) \cdot (\text{J}52\_\text{k}1 \cdot [c\_\text{Raf}] \cdot [\text{Ras}\_\text{GTP}] - \text{J}52\_\text{k}2 \cdot [c\_\text{Raf}\_\text{Ras}\_\text{GTP}])$$
 (102)

Table 160: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant       |
|--------|------|----------------|----------------|
| J52_k1 |      | 60.0           | $\overline{Z}$ |
| J52_k2 |      | 0.5            |                |

## **5.52 Reaction** J53

This is a reversible reaction of two reactants forming one product.

Name binding\_B\_Raf\_to\_Rap1\_GTP

## **Reaction equation**

$$Rap1\_GTP + B\_Raf \Longrightarrow B\_Raf\_Rap1\_GTP$$
 (103)

Table 161: Properties of each reactant.

| Id                | Name | SBO |
|-------------------|------|-----|
| Rap1_GTP<br>B_Raf |      |     |

Table 162: Properties of each product.

| Id             | Name | SBO |
|----------------|------|-----|
| B_Raf_Rap1_GTP |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{52} = vol(c1) \cdot (J53\_k1 \cdot [Rap1\_GTP] \cdot [B\_Raf] - J53\_k2 \cdot [B\_Raf\_Rap1\_GTP]) \quad (104)$$

Table 163: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant                   |
|--------|------|----------------|----------------------------|
| J53_k1 |      | 60.0           | $ \overline{\mathscr{A}} $ |
| J53_k2 |      | 0.5            |                            |

# **5.53 Reaction** J54

This is a reversible reaction of two reactants forming one product.

Name binding\_B\_Raf\_to\_Ras\_GTP

## **Reaction equation**

$$Ras\_GTP + B\_Raf \Longrightarrow B\_Raf\_Ras\_GTP$$
 (105)

### **Reactants**

Table 164: Properties of each reactant.

| Id      | Name | SBO |
|---------|------|-----|
| Ras_GTP |      |     |
| $B_Raf$ |      |     |

Table 165: Properties of each product.

| Id            | Name | SBO |
|---------------|------|-----|
| B_Raf_Ras_GTP |      |     |

**Derived unit** contains undeclared units

$$v_{53} = \text{vol}(c1) \cdot (\text{J}54\_\text{k}1 \cdot [\text{Ras\_GTP}] \cdot [\text{B\_Raf}] - \text{J}54\_\text{k}2 \cdot [\text{B\_Raf\_Ras\_GTP}])$$
 (106)

Table 166: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant               |
|--------|------|----------------|------------------------|
| J54_k1 |      | 60.0           | $ \overline{\square} $ |
| J54_k2 |      | 0.5            | $\square$              |

### 5.54 Reaction J57

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

Name ppMEK\_dephosphorylation

# **Reaction equation**

$$ppMEK \xrightarrow{PP2A} pMEK \tag{107}$$

### Reactant

Table 167: Properties of each reactant.

| Id    | Name | SBO |
|-------|------|-----|
| ррМЕК |      |     |

#### **Modifier**

Table 168: Properties of each modifier.

| Id   | Name | SBO |
|------|------|-----|
| PP2A |      |     |

Table 169: Properties of each product.

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{54} = \frac{\text{J57\_Vmax} \cdot [\text{ppMEK}] \cdot [\text{PP2A}]}{\text{J57\_Km1} + [\text{ppMEK}]}$$
(108)

Table 170: Properties of each parameter.

| Id               | Name | SBO Value Unit | Constant     |
|------------------|------|----------------|--------------|
| $J57_{Vmax}$     |      | 3.000          |              |
| ${\tt J57\_Km1}$ |      | 15.657         | $\checkmark$ |

## 5.55 Reaction J58

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

Name pMEK\_dephosphorylation

### **Reaction equation**

$$pMEK \xrightarrow{PP2A} MEK \tag{109}$$

## Reactant

Table 171: Properties of each reactant.

| Id   | Name | SBO |
|------|------|-----|
| pMEK |      |     |

#### **Modifier**

Table 172: Properties of each modifier.

| Id   | Name | SBO |
|------|------|-----|
| PP2A |      |     |

Table 173: Properties of each product.

| Id  | Name | SBO |
|-----|------|-----|
| MEK |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{55} = \text{vol}(c1) \cdot \frac{\text{J58\_Vmax} \cdot [\text{pMEK}] \cdot [\text{PP2A}]}{\text{J58\_Km1} + [\text{pMEK}]}$$
(110)

Table 174: Properties of each parameter.

| Id       | Name | SBO Value Unit | Constant  |
|----------|------|----------------|-----------|
| J58_Vmax |      | 3.000          | $\square$ |
| J58_Km1  |      | 15.657         |           |

#### **5.56 Reaction** J61

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

# Name ppMEK\_ERK

# **Reaction equation**

$$ppMEK\_ERK \xrightarrow{PP2A} pMEK\_ERK$$
 (111)

Table 175: Properties of each reactant.

| Id        | Name | SBO |
|-----------|------|-----|
| ppMEK_ERK |      |     |

#### Modifier

Table 176: Properties of each modifier.

| Id   | Name | SBO |
|------|------|-----|
| PP2A |      |     |

### **Product**

Table 177: Properties of each product.

| Id       | Name | SBO |
|----------|------|-----|
| pMEK_ERK |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{56} = \text{vol}(c1) \cdot \frac{\text{J61\_Vmax} \cdot [\text{ppMEK\_ERK}] \cdot [\text{PP2A}]}{\text{J61\_Km1} + [\text{ppMEK\_ERK}]}$$
(112)

Table 178: Properties of each parameter.

| Id                     | Name | SBO | Value  | Unit | Constant                 |
|------------------------|------|-----|--------|------|--------------------------|
| $_{	t J61\_{	t Vmax}}$ |      |     | 3.000  |      | $\overline{Z}$           |
| ${\tt J61\_Km1}$       |      |     | 15.657 |      | $\overline{\mathscr{L}}$ |

### **5.57 Reaction** J62

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

Name pMEK\_ERK\_dephosphorylation

## **Reaction equation**

$$pMEK\_ERK \xrightarrow{PP2A} MEK\_ERK$$
 (113)

Table 179: Properties of each reactant.

| Id       | Name | SBO |
|----------|------|-----|
| pMEK_ERK |      |     |

### **Modifier**

Table 180: Properties of each modifier.

| Id   | Name | SBO |
|------|------|-----|
| PP2A |      |     |

### **Product**

Table 181: Properties of each product.

| Id      | Name | SBO |
|---------|------|-----|
| MEK_ERK |      |     |

### **Kinetic Law**

Derived unit contains undeclared units

$$v_{57} = \text{vol}(c1) \cdot \frac{\text{J62\_Vmax} \cdot [\text{pMEK\_ERK}] \cdot [\text{PP2A}]}{\text{J62\_Km1} + [\text{pMEK\_ERK}]}$$
(114)

Table 182: Properties of each parameter.

| - T 1    | 3.7  | - CD-O | ***    | TT 1: |                |
|----------|------|--------|--------|-------|----------------|
| Id       | Name | SBO    | Value  | Unit  | Constant       |
| J62_Vmax |      |        | 3.000  |       | $\overline{Z}$ |
| J62_Km1  |      |        | 15.657 |       |                |

### 5.58 Reaction J63

This is a reversible reaction of one reactant forming one product.

**Name** ppERK\_dimerization

## **Reaction equation**

$$2ppERK \Longrightarrow dppERK \tag{115}$$

### Reactant

Table 183: Properties of each reactant.

| Id    | Name | SBO |
|-------|------|-----|
| ppERK |      |     |

### **Product**

Table 184: Properties of each product.

| Id     | Name | SBO |
|--------|------|-----|
| dppERK |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{58} = \text{vol}(c1) \cdot (\text{J}63\_\text{k}1 \cdot [\text{ppERK}] \cdot [\text{ppERK}] - \text{J}63\_\text{k}2 \cdot [\text{dppERK}])$$
(116)

Table 185: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant     |
|--------|------|----------------|--------------|
| J63_k1 |      | 10.000         |              |
| J63_k2 |      | 0.075          | $\checkmark$ |

## **5.59 Reaction** J66

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

Name Ras\_GTP\_dephosphorylation

## **Reaction equation**

$$Ras\_GTP \longrightarrow Ras\_GDP \tag{117}$$

Table 186: Properties of each reactant.

| Id      | Name | SBO |
|---------|------|-----|
| Ras_GTP |      |     |

Table 187: Properties of each product.

| Id      | Name | SBO |
|---------|------|-----|
| Ras_GDP |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{59} = \text{vol}(c1) \cdot \text{J66\_k} \cdot [\text{Ras\_GTP}]$$
 (118)

Table 188: Properties of each parameter.

| Id    | Name | SBO | Value                 | Unit | Constant |
|-------|------|-----|-----------------------|------|----------|
| J66_k |      | 1   | $1.667 \cdot 10^{-4}$ |      |          |

## **5.60 Reaction** J67

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

Name Rap1\_GTP\_dephosphorylation

## **Reaction equation**

$$Rap1\_GTP \longrightarrow Rap1\_GDP \tag{119}$$

Table 189: Properties of each reactant.

| Id       | Name | SBO |
|----------|------|-----|
| Rap1_GTP |      |     |

Table 190: Properties of each product.

| Id       | Name | SBO |
|----------|------|-----|
| Rap1_GDP |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{60} = \text{vol}(c1) \cdot \text{J67\_k} \cdot [\text{Rap1\_GTP}]$$
 (120)

Table 191: Properties of each parameter.

| Id    | Name | SBO | Value                 | Unit | Constant |
|-------|------|-----|-----------------------|------|----------|
| J67_k |      |     | $1.166 \cdot 10^{-4}$ | ļ    |          |

### **5.61 Reaction** J68

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

Name Rap1\_GTP\_phosphorylation

## **Reaction equation**

$$Rap1\_GDP \xrightarrow{Crk\_C3G\_pFRS2\_dpEGFR\_c\_Cbl, Crk\_C3G\_pFRS2\_dpEGFR, Crk\_C3G\_pFRS2\_pTrkA\_endo} Rap1\_GT \xrightarrow{(121)}$$

### Reactant

Table 192: Properties of each reactant.

| Id       | Name | SBO |
|----------|------|-----|
| Rap1_GDP |      |     |

### **Modifiers**

Table 193: Properties of each modifier.

| Tuble 175. I Toperties of euc  | ii iiiouiiic | 1.  |
|--|--------------|-----|
| Id   | Name         | SBO |
| Crk_C3G_pFRS2_dpEGFR_c_Cbl<br>Crk_C3G_pFRS2_dpEGFR<br>Crk_C3G_pFRS2_pTrkA_endo |              |     |

Table 194: Properties of each product.

| Id       | Name | SBO |
|----------|------|-----|
| Rap1_GTP |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$\nu_{61} = \text{vol}(\texttt{c1}) \tag{122} \\ \cdot \underbrace{ \text{J68\_Vmax} \cdot [\text{Rap1\_GDP}] \cdot ([\text{Crk\_C3G\_pFRS2\_dpEGFR}] + [\text{Crk\_C3G\_pFRS2\_dpEGFR\_c\_Cbl}] + [\text{Crk\_C3G\_pFR}] +$$

Table 195: Properties of each parameter.

| Id       | Name | SBO Value Unit | Constant                 |
|----------|------|----------------|--------------------------|
| J68_Vmax |      | 0.024          | $ \mathcal{I} $          |
| J68_Km1  |      | 0.010          | $\overline{\mathscr{L}}$ |

# **5.62 Reaction** J69

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

Name Ras\_GDP\_phosphorylation

### **Reaction equation**

$$Ras\_GDP \xrightarrow{Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl, Grb2\_SOS\_pShc\_dpEGFR, Grb2\_SOS\_pShc\_pTrkA} Ras\_GTP \xrightarrow{(123)}$$

Table 196: Properties of each reactant.

| Id      | Name | SBO |
|---------|------|-----|
| Ras_GDP |      |     |

### **Modifiers**

Table 197: Properties of each modifier.

| Tuble 1571 Troperties of cuer                      | i iiio diiiio |     |
|--|---------------|-----|
| Id   | Name          | SBO |
| Grb2_SOS_pShc_dpEGFR_c_Cbl<br>Grb2_SOS_pShc_dpEGFR |               |     |
| Grb2_SOS_pShc_pTrkA                                |               |     |

#### **Product**

Table 198: Properties of each product.

| Id      | Name | SBO |
|---------|------|-----|
| Ras_GTP |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{62} = \text{vol}(\text{c1}) \tag{124} \\ \cdot \underbrace{ \text{J69\_Vmax} \cdot [\text{Ras\_GDP}] \cdot ([\text{Grb2\_SOS\_pShc\_dpEGFR}] + [\text{Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl}] + [\text{Grb2\_SOS\_dpShc\_dpEGFR\_c\_Cbl}] + [\text{Grb2\_SOS\_dpShc\_dpEGFR\_c\_Cbl}] + [\text{Grb2\_SOS\_dpShc\_dpEGFR\_c\_Cbl}] + [\text{Grb2\_SOS\_dpShc\_dpEGFR\_c\_Cbl}] + [\text{Grb2\_SOS\_dpShc\_dpEGFR\_c\_Cbl}] + [\text{Grb2\_SOS\_dpShc\_dpEGFR\_c\_Cbl}] + [\text{Grb2\_SOS\_dpShc\_dpShc\_dpEGFR\_c\_Cbl}] + [\text{Grb2\_SOS\_dpShc\_dpEGFR\_c\_Cbl}] + [\text{Grb2\_SOS\_dpShc\_dpEGFR\_c\_Cbl}] + [\text{Grb2\_SOS\_dpShc\_dpShc\_dpEGFR\_c\_Cbl}] + [\text{Grb2\_SOS\_dpShc\_d$$

Table 199: Properties of each parameter.

| Id       | Name | SBO Value Unit | Constant  |
|----------|------|----------------|-----------|
| J69_Vmax |      | 2.00           | $\square$ |
| J69_Km1  |      | 0.02           | $\square$ |

# 5.63 Reaction J70

This is a reversible reaction of two reactants forming one product.

Name binding\_NGF\_to\_NGFR

# **Reaction equation**

$$NGF + NGFR \rightleftharpoons L\_NGFR$$
 (125)

#### **Reactants**

Table 200: Properties of each reactant.

| Id   | Name | SBO |
|------|------|-----|
| NGF  |      |     |
| NGFR |      |     |

#### **Product**

Table 201: Properties of each product.

| Id     | Name | SBO |
|--------|------|-----|
| L_NGFR |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{63} = \text{vol}\left(\text{compartment}\right) \cdot \left(\text{J70\_k1} \cdot \left[\text{NGF}\right] \cdot \left[\text{NGFR}\right] - \text{J70\_k2} \cdot \left[\text{L\_NGFR}\right]\right)$$
 (126)

Table 202: Properties of each parameter.

| Id     | Name | SBO | Value             | Unit | Constant                |
|--------|------|-----|-------------------|------|-------------------------|
| J70_k1 |      |     | 6.200             |      | $\blacksquare$          |
| J70_k2 |      |     | $6.4\cdot10^{-5}$ |      | $\overline{\mathbf{Z}}$ |

### 5.64 Reaction J71

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

Name TrkA\_phosphorylation

# **Reaction equation**

$$L\_NGFR \longrightarrow pTrkA$$
 (127)

#### Reactant

Table 203: Properties of each reactant.

| Id          | Name | SBO |
|-------------|------|-----|
| $L_{-}NGFR$ |      |     |

### **Product**

Table 204: Properties of each product.

| Id    | Name | SBO |
|-------|------|-----|
| pTrkA |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{64} = \text{vol} \left( \text{compartment} \right) \cdot \text{J71\_k} \cdot \left[ \text{L\_NGFR} \right]$$
 (128)

Table 205: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant     |
|-------|------|----------------|--------------|
| J71_k |      | 1.0            | $\checkmark$ |

### **5.65 Reaction** J72

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

Name pTrkA\_intermalization

# **Reaction equation**

$$pTrkA \longrightarrow pTrkA\_endo$$
 (129)

#### Reactant

Table 206: Properties of each reactant.

| Id    | Name | SBO |
|-------|------|-----|
| pTrkA |      |     |

### **Product**

Table 207: Properties of each product.

| Id         | Name | SBO |
|------------|------|-----|
| pTrkA_endo |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{65} = \text{vol}(c1) \cdot \text{J72\_k} \cdot [\text{pTrkA}] \tag{130}$$

Table 208: Properties of each parameter.

| Id    | Name | SBO | Value             | Unit | Constant |
|-------|------|-----|-------------------|------|----------|
| J72_k |      |     | $6.3\cdot10^{-4}$ |      |          |

### 5.66 Reaction J73

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

Name pTrkA\_endo\_degradation

# **Reaction equation**

$$pTrkA\_endo \longrightarrow degradation$$
 (131)

### Reactant

Table 209: Properties of each reactant.

| Id         | Name | SBO |
|------------|------|-----|
| pTrkA_endo |      |     |

### **Product**

Table 210: Properties of each product.

| Id          | Name | SBO |
|-------------|------|-----|
| degradation |      |     |

|--|

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{66} = \text{vol}(c1) \cdot \text{J73\_k} \cdot [\text{pTrkA\_endo}]$$
 (132)

Table 211: Properties of each parameter.

| Id    | Name | SBO | Value             | Unit | Constant |
|-------|------|-----|-------------------|------|----------|
| J73_k |      |     | $4.2\cdot10^{-4}$ |      |          |

# **5.67 Reaction** J74

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

Name pTrkA\_degradation

# **Reaction equation**

$$pTrkA \longrightarrow degradation$$
 (133)

# Reactant

Table 212: Properties of each reactant.

| Id    | Name | SBO |
|-------|------|-----|
| pTrkA |      |     |

### **Product**

Table 213: Properties of each product.

| Id          | Name | SBO |
|-------------|------|-----|
| degradation |      |     |

#### **Kinetic Law**

$$v_{67} = \text{vol}(c1) \cdot \text{J74\_k} \cdot [\text{pTrkA}] \tag{134}$$

Table 214: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant       |
|-------|------|----------------|----------------|
| J74_k |      | 0.002          | $\overline{Z}$ |

### **5.68 Reaction** J75

This is a reversible reaction of two reactants forming one product.

Name binding\_Shc\_to\_pTrkA

# **Reaction equation**

$$Shc + pTrkA \Longrightarrow Shc - pTrkA \tag{135}$$

#### **Reactants**

Table 215: Properties of each reactant.

| Id    | Name | SBO |
|-------|------|-----|
| Shc   |      |     |
| pTrkA |      |     |

#### **Product**

Table 216: Properties of each product.

| Id        | Name | SBO |
|-----------|------|-----|
| Shc_pTrkA |      |     |

#### **Kinetic Law**

$$v_{68} = \text{vol}(c1) \cdot (J75 \underline{k}1 \cdot [Shc] \cdot [pTrkA] - J75 \underline{k}2 \cdot [Shc \underline{p}TrkA])$$
(136)

Table 217: Properties of each parameter.

|        |      |           | •      |           |
|--------|------|-----------|--------|-----------|
| Id     | Name | SBO Value | e Unit | Constant  |
| J75_k1 |      | 10.0      |        | $\square$ |
| J75_k2 |      | 0.2       |        |           |

### **5.69 Reaction** J76

This is a reversible reaction of two reactants forming one product.

Name binding\_pShc\_to\_pTrkA

# **Reaction equation**

$$pShc + pTrkA \Longrightarrow pShc_pTrkA$$
 (137)

#### **Reactants**

Table 218: Properties of each reactant.

| Id            | Name | SBO |
|---------------|------|-----|
| pShc<br>pTrkA |      |     |

#### **Product**

Table 219: Properties of each product.

| Id         | Name | SBO |
|------------|------|-----|
| pShc_pTrkA |      |     |

#### **Kinetic Law**

$$v_{69} = \text{vol}(c1) \cdot (J76 \underline{k} 1 \cdot [pShc] \cdot [pTrkA] - J76 \underline{k} 2 \cdot [pShc \underline{pTrkA}])$$
(138)

Table 220: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J76_k1 |      | 10.0           |          |
| J76_k2 |      | 0.2            |          |

### 5.70 Reaction J77

This is a reversible reaction of two reactants forming one product.

Name binding\_FRS2\_to\_pTrkA

# **Reaction equation**

$$FRS2 + pTrkA \Longrightarrow FRS2 - pTrkA \tag{139}$$

### **Reactants**

Table 221: Properties of each reactant.

| Id    | Name | SBO |
|-------|------|-----|
| FRS2  |      |     |
| pTrkA |      |     |

### **Product**

Table 222: Properties of each product.

| Id         | Name | SBO |
|------------|------|-----|
| FRS2_pTrkA |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{70} = \text{vol}(c1) \cdot (J77 k1 \cdot [FRS2] \cdot [pTrkA] - J77 k2 \cdot [FRS2 pTrkA])$$

$$(140)$$

Table 223: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant  |
|--------|------|----------------|-----------|
| J77_k1 |      | 5.0            | $\square$ |
| J77_k2 |      | 0.1            | $\square$ |

### 5.71 Reaction J78

This is a reversible reaction of two reactants forming one product.

Name binding\_pFRS2\_to\_pTrkA

# **Reaction equation**

$$pFRS2 + pTrkA \Longrightarrow pFRS2 pTrkA$$
 (141)

#### **Reactants**

Table 224: Properties of each reactant.

| Id    | Name | SBO |
|-------|------|-----|
| pFRS2 |      |     |
| pTrkA |      |     |

#### **Product**

Table 225: Properties of each product.

| Id          | Name | SBO |
|-------------|------|-----|
| pFRS2_pTrkA |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{71} = vol(c1) \cdot (J78\_k1 \cdot [pFRS2] \cdot [pTrkA] - J78\_k2 \cdot [pFRS2\_pTrkA])$$
 (142)

Table 226: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant  |
|--------|------|----------------|-----------|
| J78_k1 |      | 5.0            |           |
| J78_k2 |      | 0.1            | $\square$ |

### 5.72 Reaction J79

This is a reversible reaction of two reactants forming one product.

Name binding\_Shc\_to\_pTrkA\_endo

# **Reaction equation**

$$pTrkA\_endo + Shc \Longrightarrow Shc\_pTrkA\_endo$$
 (143)

#### **Reactants**

Table 227: Properties of each reactant.

| Id                | Name | SBO |
|-------------------|------|-----|
| pTrkA_endo<br>Shc |      |     |

#### **Product**

Table 228: Properties of each product.

| Id             | Name | SBO |
|----------------|------|-----|
| Shc_pTrkA_endo |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{72} = \text{vol}(c1) \cdot (J79 k1 \cdot [pTrkA\_endo] \cdot [Shc] - J79 k2 \cdot [Shc\_pTrkA\_endo])$$
 (144)

Table 229: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant       |
|--------|------|----------------|----------------|
| J79_k1 |      | 10.0           | $\overline{Z}$ |
| J79_k2 |      | 0.2            | $\square$      |

### 5.73 Reaction J80

This is a reversible reaction of two reactants forming one product.

 $\textbf{Name} \ binding\_pShc\_to\_pTrkA\_endo$ 

# **Reaction equation**

$$pTrkA\_endo + pShc \Longrightarrow pShc\_pTrkA\_endo$$
 (145)

### **Reactants**

Table 230: Properties of each reactant.

| Id                 | Name | SBO |
|--------------------|------|-----|
| pTrkA_endo<br>pShc |      |     |

### **Product**

Table 231: Properties of each product.

| Id              | Name | SBO |
|-----------------|------|-----|
| pShc_pTrkA_endo |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{73} = \text{vol}(c1) \cdot (\text{J}80\_\text{k}1 \cdot [\text{pTrkA\_endo}] \cdot [\text{pShc}] - \text{J}80\_\text{k}2 \cdot [\text{pShc\_pTrkA\_endo}])$$
 (146)

Table 232: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant                  |
|--------|------|----------------|---------------------------|
| J80_k1 |      | 10.0           | $\overline{\hspace{1cm}}$ |
| J80_k2 |      | 0.2            |                           |

# 5.74 Reaction J81

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

Name Shc\_pTrkA\_endo\_phosphorylation

# **Reaction equation**

$$Shc_pTrkA\_endo \longrightarrow pShc\_pTrkA\_endo$$
 (147)

# Reactant

Table 233: Properties of each reactant.

| Id             | Name | SBO |
|----------------|------|-----|
| Shc_pTrkA_endo |      |     |

### **Product**

Table 234: Properties of each product.

| Id              | Name |  |
|-----------------|------|--|
| pShc_pTrkA_endo |      |  |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{74} = \text{vol}(c1) \cdot \text{J81\_k} \cdot [\text{Shc\_pTrkA\_endo}]$$
 (148)

Table 235: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J81_k |      | 0.1            |          |

### 5.75 Reaction J82

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

Name Shc\_pTrkA\_phosphorylation

# **Reaction equation**

$$Shc_pTrkA \longrightarrow pShc_pTrkA$$
 (149)

#### Reactant

Table 236: Properties of each reactant.

| Id        | Name | SBO |
|-----------|------|-----|
| Shc_pTrkA |      |     |

### **Product**

Table 237: Properties of each product.

| Id         | Name | SBO |
|------------|------|-----|
| pShc_pTrkA |      |     |

| Id | Name | SBO |
|----|------|-----|
| -  |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{75} = \text{vol}(c1) \cdot \text{J82\_k} \cdot [\text{Shc\_pTrkA}]$$
 (150)

Table 238: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J82_k |      | 0.1            |          |

# 5.76 Reaction J83

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

Name pFRS2\_pTrkA\_phosphorylation

# **Reaction equation**

$$FRS2\_pTrkA \longrightarrow pFRS2\_pTrkA \tag{151}$$

### Reactant

Table 239: Properties of each reactant.

| Id         | Name | SBO |
|------------|------|-----|
| FRS2_pTrkA |      |     |

### **Product**

Table 240: Properties of each product.

| Id          | Name | SBO |
|-------------|------|-----|
| pFRS2_pTrkA |      |     |

#### **Kinetic Law**

$$v_{76} = \text{vol}(c1) \cdot \text{J83\_k} \cdot [\text{FRS2\_pTrkA}]$$
 (152)

Table 241: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J83_k |      | 2.0            |          |

### 5.77 Reaction J84

This is a reversible reaction of two reactants forming one product.

Name binding\_FRS2\_to\_pTrkA\_endo

# **Reaction equation**

$$pTrkA\_endo + FRS2 \Longrightarrow FRS2\_pTrkA\_endo$$
 (153)

#### **Reactants**

Table 242: Properties of each reactant.

| Id                 | Name | SBO |
|--------------------|------|-----|
| pTrkA_endo<br>FRS2 |      |     |

#### **Product**

Table 243: Properties of each product.

| 1               | 1    |     |
|-----------------|------|-----|
| Id              | Name | SBO |
| FRS2_pTrkA_endo |      |     |

#### **Kinetic Law**

$$v_{77} = \text{vol}(c1) \cdot (J84 \text{\_k}1 \cdot [pTrkA\_endo] \cdot [FRS2] - J84 \text{\_k}2 \cdot [FRS2\_pTrkA\_endo])$$
 (154)

Table 244: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J84_k1 |      | 5.0            |          |
| J84_k2 |      | 0.1            |          |

### 5.78 Reaction J85

This is a reversible reaction of two reactants forming one product.

Name binding\_pFRS2\_to\_pTrkA\_endo

### **Reaction equation**

$$pTrkA\_endo + pFRS2 \Longrightarrow pFRS2\_pTrkA\_endo$$
 (155)

#### **Reactants**

Table 245: Properties of each reactant.

| Id         | Name | SBO |
|------------|------|-----|
| pTrkA_endo |      |     |
| pFRS2      |      |     |

#### **Product**

Table 246: Properties of each product.

| Id               | Name | SBO |
|------------------|------|-----|
| pFRS2_pTrkA_endo |      |     |

#### **Kinetic Law**

$$v_{78} = \text{vol}(c1) \cdot (J85 \underline{k}1 \cdot [pTrkA\_endo] \cdot [pFRS2] - J85 \underline{k}2 \cdot [pFRS2\_pTrkA\_endo])$$
 (156)

Table 247: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant  |
|--------|------|----------------|-----------|
| J85_k1 |      | 5.0            |           |
| J85_k2 |      | 0.1            | $\square$ |

### 5.79 Reaction J86

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

Name FRS2\_pTrkA\_endo\_phosphorylation

# **Reaction equation**

$$FRS2\_pTrkA\_endo \longrightarrow pFRS2\_pTrkA\_endo$$
 (157)

### Reactant

Table 248: Properties of each reactant.

Id Name SBO

FRS2\_pTrkA\_endo

### **Product**

Table 249: Properties of each product.

| Id               | Name | SBO |
|------------------|------|-----|
| pFRS2_pTrkA_endo |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{79} = \text{vol}(c1) \cdot \text{J86\_k} \cdot [\text{FRS2\_pTrkA\_endo}]$$
 (158)

Table 250: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J86_k |      | 2.0            |          |

### 5.80 Reaction J87

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

Name FRS2\_pTrkA\_degradation

### **Reaction equation**

$$FRS2\_pTrkA \longrightarrow degradation + FRS2$$
 (159)

### Reactant

Table 251: Properties of each reactant.

| Id         | Name | SBO |
|------------|------|-----|
| FRS2_pTrkA |      |     |

### **Products**

Table 252: Properties of each product.

| Id                  | Name | SBO |
|---------------------|------|-----|
| degradation<br>FRS2 |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{80} = \text{vol}(c1) \cdot \text{J87\_k} \cdot [\text{FRS2\_pTrkA}]$$
 (160)

Table 253: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant  |
|-------|------|----------------|-----------|
| J87_k |      | 0.002          | $\square$ |

### **5.81 Reaction** J88

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

Name pFRS2\_pTrkA\_degradation

# **Reaction equation**

$$pFRS2_pTrkA \longrightarrow degradation + pFRS2$$
 (161)

# Reactant

Table 254: Properties of each reactant.

| Id          | Name | SBO |
|-------------|------|-----|
| pFRS2_pTrkA |      |     |

### **Products**

Table 255: Properties of each product.

| Name | SBO  |
|------|------|
|      |      |
|      | Name |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{81} = \text{vol}(c1) \cdot J88 \underline{k} \cdot [pFRS2 \underline{p}TrkA]$$
 (162)

Table 256: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant  |
|-------|------|----------------|-----------|
| J88_k |      | 0.002          | $\square$ |

# 5.82 Reaction J89

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

Name Shc\_pTrkA\_degradation

# **Reaction equation**

$$Shc_pTrkA \longrightarrow degradation + Shc$$
 (163)

### Reactant

Table 257: Properties of each reactant.

| Id        | Name | SBO |
|-----------|------|-----|
| Shc_pTrkA |      |     |

### **Products**

Table 258: Properties of each product.

| Id                 | Name | SBO |
|--------------------|------|-----|
| degradation<br>Shc |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{82} = \text{vol}(c1) \cdot J89 \underline{k} \cdot [\text{Shc\_pTrkA}]$$
 (164)

Table 259: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant |
|-------|------|----------------|----------|
| J89_k |      | 0.002          |          |

# **5.83 Reaction** J90

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

Name pShc\_pTrkA\_degradation

# **Reaction equation**

$$pShc_pTrkA \longrightarrow degradation + pShc$$
 (165)

### Reactant

Table 260: Properties of each reactant.

| Id         | Name | SBO |
|------------|------|-----|
| pShc_pTrkA |      |     |

### **Products**

Table 261: Properties of each product.

| Id               | Name | SBO |
|------------------|------|-----|
| degradation pShc |      |     |

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{83} = \text{vol}(c1) \cdot \text{J}90\_\text{k} \cdot [\text{pShc\_pTrkA}]$$
 (166)

Table 262: Properties of each parameter.

| Id    | Name | SBO Value Unit | Constant     |
|-------|------|----------------|--------------|
| J90_k |      | 0.002          | $\checkmark$ |

### 5.84 Reaction J92

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

Name FRS2\_pTrkA\_endo\_degradation

# **Reaction equation**

$$FRS2_pTrkA\_endo \longrightarrow degradation + FRS2$$
 (167)

#### Reactant

Table 263: Properties of each reactant.

| Id              | Name | SBO |
|-----------------|------|-----|
| FRS2_pTrkA_endo |      |     |

### **Products**

Table 264: Properties of each product.

| Id                  | Name | SBO |
|---------------------|------|-----|
| degradation<br>FRS2 |      |     |

### **Kinetic Law**

Derived unit contains undeclared units

$$v_{84} = \text{vol}(c1) \cdot \text{J92\_k} \cdot [\text{FRS2\_pTrkA\_endo}]$$
 (168)

Table 265: Properties of each parameter.

| Id    | Name | SBO | Value              | Unit | Constant |
|-------|------|-----|--------------------|------|----------|
| J92_k |      |     | $4.2\cdot 10^{-4}$ |      |          |

### 5.85 Reaction J93

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

Name Shc\_pTrkA\_endo\_degradation

### **Reaction equation**

$$Shc_pTrkA\_endo \longrightarrow degradation + Shc$$
 (169)

#### Reactant

Table 266: Properties of each reactant.

| Id             | Name | SBO |
|----------------|------|-----|
| Shc_pTrkA_endo |      |     |

#### **Products**

Table 267: Properties of each product.

| Id                 | Name | SBO |
|--------------------|------|-----|
| degradation<br>Shc |      |     |

# **Kinetic Law**

$$v_{85} = \text{vol}(\text{c1}) \cdot \text{J93\_k} \cdot [\text{Shc\_pTrkA\_endo}]$$
 (170)

Table 268: Properties of each parameter.

| Id    | Name | SBO | Value             | Unit | Constant |
|-------|------|-----|-------------------|------|----------|
| J93_k |      |     | $4.2\cdot10^{-4}$ |      |          |

### 5.86 Reaction J94

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

Name pShc\_pTrkA\_endo\_degradation

# **Reaction equation**

$$pShc\_pTrkA\_endo \longrightarrow degradation + pShc$$
 (171)

#### Reactant

Table 269: Properties of each reactant.

| Id              | Name | SBO |
|-----------------|------|-----|
| pShc_pTrkA_endo |      |     |

### **Products**

Table 270: Properties of each product.

| Id               | Name | SBO |
|------------------|------|-----|
| degradation pShc |      |     |

# **Kinetic Law**

$$v_{86} = \text{vol}(c1) \cdot \text{J94\_k} \cdot [\text{pShc\_pTrkA\_endo}]$$
 (172)

Table 271: Properties of each parameter.

| Id    | Name | SBO | Value              | Unit | Constant |
|-------|------|-----|--------------------|------|----------|
| J94_k |      |     | $4.2\cdot 10^{-4}$ |      |          |

### 5.87 Reaction J95

This is a reversible reaction of two reactants forming one product.

Name binding\_Grb2\_SOS\_to\_pShc\_pTrkA\_endo

# **Reaction equation**

$$SOS\_Grb2 + pShc\_pTrkA\_endo \Longrightarrow Grb2\_SOS\_pShc\_pTrkA\_endo$$
 (173)

### **Reactants**

Table 272: Properties of each reactant.

| Id                        | Name | SBO |
|---------------------------|------|-----|
| SOS_Grb2                  |      |     |
| ${\tt pShc\_pTrkA\_endo}$ |      |     |

### **Product**

Table 273: Properties of each product.

| Id                       | Name | SBO |
|--------------------------|------|-----|
| Grb2_SOS_pShc_pTrkA_endo |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{87} = \text{vol}(\text{c1}) \cdot (\text{J95\_k1} \cdot [\text{SOS\_Grb2}] \cdot [\text{pShc\_pTrkA\_endo}] - \text{J95\_k2}$$

$$\cdot [\text{Grb2\_SOS\_pShc\_pTrkA\_endo}])$$
(174)

Table 274: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J95_k1 |      | 10.0           |          |
| J95_k2 |      | 0.2            |          |

### 5.88 Reaction J96

This is a reversible reaction of two reactants forming one product.

Name binding\_Grb2\_SOS\_to\_pShc\_pTrkA

# **Reaction equation**

$$SOS\_Grb2 + pShc\_pTrkA \Longrightarrow Grb2\_SOS\_pShc\_pTrkA$$
 (175)

#### **Reactants**

Table 275: Properties of each reactant.

| Id         | Name | SBO |
|------------|------|-----|
| SOS_Grb2   |      |     |
| pShc_pTrkA |      |     |

#### **Product**

Table 276: Properties of each product.

| Id                  | Name | SBO |
|---------------------|------|-----|
| Grb2_SOS_pShc_pTrkA |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{88} = \text{vol}(c1) \cdot (\text{J96\_k1} \cdot [\text{SOS\_Grb2}] \cdot [\text{pShc\_pTrkA}] - \text{J96\_k2} \cdot [\text{Grb2\_SOS\_pShc\_pTrkA}]) \quad (176)$$

Table 277: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J96_k1 |      | 10.0           |          |
| J96_k2 |      | 0.2            |          |

### 5.89 Reaction J97

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

Name Grb2\_SOS\_pShc\_pTrkA\_ubiquitination

# **Reaction equation**

$$Grb2\_SOS\_pShc\_pTrkA \longrightarrow Grb2\_SOS\_pShc\_pTrkA\_endo$$
 (177)

#### Reactant

Table 278: Properties of each reactant.

| Table 270. Troperties of | cacii ica | ctant. |
|--------------------------|-----------|--------|
| Id                       | Name      | SBO    |
| Grb2_SOS_pShc_pTrkA      |           |        |

### **Product**

Table 279: Properties of each product.

| Id                       | Name | SBO |
|--------------------------|------|-----|
| Grb2_SOS_pShc_pTrkA_endo |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{89} = \text{vol}(c1) \cdot \text{J97\_k} \cdot [\text{Grb2\_SOS\_pShc\_pTrkA}]$$
 (178)

Table 280: Properties of each parameter.

| Id    | Name | SBO | Value              | Unit | Constant |
|-------|------|-----|--------------------|------|----------|
| J97_k |      |     | $6.3\cdot 10^{-4}$ |      |          |

### **5.90 Reaction** J98

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

Name Crk\_C3G\_pFRS2\_pTrkA\_ubiquitination

# **Reaction equation**

$$Crk_C3G_pFRS2_pTrkA \longrightarrow Crk_C3G_pFRS2_pTrkA_endo$$
 (179)

### Reactant

Table 281: Properties of each reactant.

| Id                  | Name | SBO |
|---------------------|------|-----|
| Crk_C3G_pFRS2_pTrkA |      |     |

### **Product**

Table 282: Properties of each product.

| ruote 202: I roperties of eu | en produc | <del></del> |
|------------------------------|-----------|-------------|
| Id                           | Name      | SBO         |
| Crk_C3G_pFRS2_pTrkA_endo     |           |             |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{90} = \text{vol}(c1) \cdot \text{J98\_k} \cdot [\text{Crk\_C3G\_pFRS2\_pTrkA}]$$
(180)

Table 283: Properties of each parameter.

| Id    | Name | SBO | Value              | Unit | Constant |
|-------|------|-----|--------------------|------|----------|
| J98_k |      |     | $6.3\cdot 10^{-4}$ |      |          |

### 5.91 Reaction J99

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

Name pFRS2\_pTrkA\_ubiquitination

# **Reaction equation**

$$pFRS2\_pTrkA \longrightarrow pFRS2\_pTrkA\_endo$$
 (181)

#### Reactant

Table 284: Properties of each reactant.

| Id          | Name | SBO |
|-------------|------|-----|
| pFRS2_pTrkA |      |     |

### **Product**

Table 285: Properties of each product.

|                  | 1    |     |
|------------------|------|-----|
| Id               | Name | SBO |
| pFRS2_pTrkA_endo |      |     |

|  | Id | Name | SBO |
|--|----|------|-----|
|--|----|------|-----|

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{91} = \text{vol}(c1) \cdot \text{J99\_k} \cdot [\text{pFRS2\_pTrkA}]$$
 (182)

Table 286: Properties of each parameter.

| Id    | Name | SBO | Value             | Unit | Constant |
|-------|------|-----|-------------------|------|----------|
| J99_k |      |     | $6.3\cdot10^{-4}$ |      |          |

### **5.92 Reaction** J100

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

Name FRS2\_pTrkA\_ubiquitination

# **Reaction equation**

$$FRS2\_pTrkA \longrightarrow FRS2\_pTrkA\_endo$$
 (183)

### Reactant

Table 287: Properties of each reactant.

| Id         | Name | SBO |
|------------|------|-----|
| FRS2_pTrkA |      |     |

# **Product**

Table 288: Properties of each product.

| Id              | Name | SBO |
|-----------------|------|-----|
| FRS2_pTrkA_endo |      |     |

#### **Kinetic Law**

$$v_{92} = vol(c1) \cdot J100 \underline{\ k} \cdot [FRS2 \underline{\ pTrkA}]$$
 (184)

Table 289: Properties of each parameter.

| Id     | Name | SBO | Value             | Unit | Constant       |
|--------|------|-----|-------------------|------|----------------|
| J100_k |      |     | $6.3\cdot10^{-4}$ |      | $\overline{Z}$ |

### **5.93 Reaction** J101

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

Name pShc\_pTrkA\_ubiquitination

# **Reaction equation**

$$pShc_pTrkA \longrightarrow pShc_pTrkA\_endo$$
 (185)

#### Reactant

Table 290: Properties of each reactant.

| Id         | Name | SBO |
|------------|------|-----|
| pShc_pTrkA |      |     |

#### **Product**

Table 291: Properties of each product.

| Id              | Name | SBO |
|-----------------|------|-----|
| pShc_pTrkA_endo |      |     |

#### **Kinetic Law**

$$v_{93} = \text{vol}(c1) \cdot \text{J}101 \underline{\text{k}} \cdot [\text{pShc\_pTrkA}]$$
 (186)

Table 292: Properties of each parameter.

| Id     | Name | SBO | Value             | Unit | Constant |
|--------|------|-----|-------------------|------|----------|
| J101_k |      |     | $6.3\cdot10^{-4}$ |      |          |

### **5.94 Reaction** J102

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

Name Shc\_pTrkA\_ubiquitination

# **Reaction equation**

$$Shc_pTrkA \longrightarrow Shc_pTrkA\_endo$$
 (187)

### Reactant

Table 293: Properties of each reactant.

| Id        | Name | SBO |
|-----------|------|-----|
| Shc_pTrkA | ·    |     |

### **Product**

Table 294: Properties of each product.

| Id             | Name | SBO |
|----------------|------|-----|
| Shc_pTrkA_endo |      |     |

# **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{94} = \text{vol}(c1) \cdot \text{J}102\_k \cdot [\text{Shc\_pTrkA}]$$
(188)

Table 295: Properties of each parameter.

| Id     | Name | SBO | Value             | Unit | Constant   |
|--------|------|-----|-------------------|------|------------|
| J102_k |      |     | $6.3\cdot10^{-4}$ |      | lacksquare |

# **5.95 Reaction** J103

This is a reversible reaction of two reactants forming one product.

Name binding\_Crk\_C3G\_to\_pFRS2\_pTrkA

### **Reaction equation**

$$Crk\_C3G + pFRS2\_pTrkA \Longrightarrow Crk\_C3G\_pFRS2\_pTrkA$$
 (189)

#### **Reactants**

Table 296: Properties of each reactant.

| Id          | Name | SBO |
|-------------|------|-----|
| Crk_C3G     |      |     |
| pFRS2_pTrkA |      |     |

#### **Product**

Table 297: Properties of each product.

| Id                  | Name | SBO |
|---------------------|------|-----|
| Crk_C3G_pFRS2_pTrkA |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{95} = \text{vol}(\text{c1}) \cdot (\text{J}103\_\text{k}1 \cdot [\text{Crk\_C3G}] \cdot [\text{pFRS2\_pTrkA}] - \text{J}103\_\text{k}2 \cdot [\text{Crk\_C3G\_pFRS2\_pTrkA}])$$

$$(190)$$

Table 298: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant                     |
|---------|------|----------------|------------------------------|
| J103_k1 |      | 1.0            | $\overline{Z}$               |
| J103_k2 |      | 0.2            | $   \overline{\mathscr{L}} $ |

### **5.96 Reaction** J104

This is a reversible reaction of two reactants forming one product.

Name binding\_Crk\_C3G\_to\_pFRS2\_pTrkA\_endo

### **Reaction equation**

$$Crk_C3G + pFRS2_pTrkA_endo \rightleftharpoons Crk_C3G_pFRS2_pTrkA_endo$$
 (191)

#### **Reactants**

Table 299: Properties of each reactant.

| Tuble 255. Troperties       | Tuble 277. I Toperties of each reactant. |     |  |  |
|-----------------------------|--|-----|--|--|
| Id                          | Name                                     | SBO |  |  |
| Crk_C3G<br>pFRS2_pTrkA_endo |  |     |  |  |

#### **Product**

Table 300: Properties of each product.

| Id                       | Name | SBO |
|--------------------------|------|-----|
| Crk_C3G_pFRS2_pTrkA_endo |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{96} = vol(c1) \cdot (J104\_k1 \cdot [Crk\_C3G] \cdot [pFRS2\_pTrkA\_endo] - J104\_k2$$

$$\cdot [Crk\_C3G\_pFRS2\_pTrkA\_endo])$$
(192)

Table 301: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant |
|---------|------|----------------|----------|
| J104_k1 |      | 1.0            |          |
| J104_k2 |      | 0.2            |          |

### **5.97 Reaction** J105

This is a reversible reaction of two reactants forming one product.

Name binding\_Grb2\_SOS\_pShc\_to\_pTrkA

### **Reaction equation**

$$Grb2\_SOS\_pShc + pTrkA \Longrightarrow Grb2\_SOS\_pShc\_pTrkA$$
 (193)

#### **Reactants**

Table 302: Properties of each reactant.

| Id            | Name | SBO |
|---------------|------|-----|
| Grb2_SOS_pShc |      |     |
| pTrkA         |      |     |

### **Product**

Table 303: Properties of each product.

| Id                  | Name | SBO |
|---------------------|------|-----|
| Grb2_SOS_pShc_pTrkA |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$\nu_{97} = vol\left(c1\right) \cdot \left(J105\_k1 \cdot \left[Grb2\_SOS\_pShc\right] \cdot \left[pTrkA\right] - J105\_k2 \cdot \left[Grb2\_SOS\_pShc\_pTrkA\right]\right) \tag{194}$$

Table 304: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant  |
|---------|------|----------------|-----------|
| J105_k1 |      | 10.0           | $\square$ |
| J105_k2 |      | 0.2            | $\square$ |

### **5.98 Reaction** J106

This is a reversible reaction of two reactants forming one product.

 $\textbf{Name} \ binding\_Grb2\_SOS\_pShc\_to\_pTrkA\_endo$ 

# **Reaction equation**

$$Grb2\_SOS\_pShc + pTrkA\_endo \Longrightarrow Grb2\_SOS\_pShc\_pTrkA\_endo$$
 (195)

# Reactants

Table 305: Properties of each reactant.

| Id                          | Name | SBO |
|-----------------------------|------|-----|
| Grb2_SOS_pShc<br>pTrkA_endo |      |     |

### **Product**

Table 306: Properties of each product.

| Id                       | Name | SBO |
|--------------------------|------|-----|
| Grb2_SOS_pShc_pTrkA_endo |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{98} = vol(c1) \cdot (J106\_k1 \cdot [Grb2\_SOS\_pShc] \cdot [pTrkA\_endo] - J106\_k2$$

$$\cdot [Grb2\_SOS\_pShc\_pTrkA\_endo])$$

$$(196)$$

Table 307: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant                |
|---------|------|----------------|-------------------------|
| J106_k1 |      | 10.0           | $\overline{Z}$          |
| J106_k2 |      | 0.2            | $\overline{\mathbf{Z}}$ |

### **5.99 Reaction** J107

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

Name Crk\_C3G\_pFRS2\_pTrkA\_degradation

# **Reaction equation**

$$Crk_C3G_pFRS2_pTrkA \longrightarrow degradation + pFRS2 + Crk_C3G$$
 (197)

### Reactant

| Table 308: Properties of each reactant. |
|---|
|---|

| Id                  | Name |  |
|---------------------|------|--|
| Crk_C3G_pFRS2_pTrkA |      |  |

### **Products**

Table 309: Properties of each product.

| Id                              | Name | SBO |
|---------------------------------|------|-----|
| degradation<br>pFRS2<br>Crk_C3G |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{99} = \text{vol}(c1) \cdot \text{J}107\_k \cdot [\text{Crk\_C3G\_pFRS2\_pTrkA}]$$
 (198)

Table 310: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J107_k |      | 0.002          |          |

# **5.100 Reaction** J108

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

 $\textbf{Name} \ \ Crk\_C3G\_pFRS2\_pTrkA\_endo\_degradation$ 

# **Reaction equation**

$$Crk\_C3G\_pFRS2\_pTrkA\_endo \longrightarrow degradation + Crk\_C3G + pFRS2$$
 (199)

### Reactant

Table 311: Properties of each reactant.

| Id                       | Name | SBO |
|--------------------------|------|-----|
| Crk_C3G_pFRS2_pTrkA_endo |      |     |

#### **Products**

Table 312: Properties of each product.

| Id          | Name | SBO |
|-------------|------|-----|
| degradation |      |     |
| $Crk_C3G$   |      |     |
| pFRS2       |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{100} = vol(c1) \cdot J108 \underline{k} \cdot [Crk\underline{C3G}pFRS2\underline{p}TrkA\underline{endo}]$$
 (200)

Table 313: Properties of each parameter.

| Id     | Name | SBO | Value             | Unit | Constant |
|--------|------|-----|-------------------|------|----------|
| J108_k |      | 4.3 | $2 \cdot 10^{-4}$ |      |          |

### **5.101 Reaction** J109

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

Name Grb2\_SOS\_pShc\_pTrkA\_degradation

### **Reaction equation**

$$Grb2\_SOS\_pShc\_pTrkA \longrightarrow degradation + Grb2\_SOS\_pShc$$
 (201)

### Reactant

Table 314: Properties of each reactant.

| Id                  | Name | SBO |
|---------------------|------|-----|
| Grb2_SOS_pShc_pTrkA |      |     |

### **Products**

Table 315: Properties of each product.

| Id                           | Name | SBO |
|------------------------------|------|-----|
| degradation<br>Grb2_SOS_pShc |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{101} = \text{vol}(c1) \cdot \text{J}109\_\text{k} \cdot [\text{Grb2\_SOS\_pShc\_pTrkA}]$$
 (202)

Table 316: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant  |
|--------|------|----------------|-----------|
| J109_k |      | 0.002          | $\square$ |

### **5.102 Reaction** J110

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

Name Grb2\_SOS\_pShc\_pTrkA\_endo\_degradation

### **Reaction equation**

$$Grb2\_SOS\_pShc\_pTrkA\_endo \longrightarrow degradation + Grb2\_SOS\_pShc$$
 (203)

#### Reactant

Table 317: Properties of each reactant.

| Tuble 317.11 openies of each reactain. |      |     |  |
|--|------|-----|--|
| Id                                     | Name | SBO |  |
| Grb2_SOS_pShc_pTrkA_endo               |      |     |  |

### **Products**

Table 318: Properties of each product.

| Id                           | Name | SBO |
|------------------------------|------|-----|
| degradation<br>Grb2_SOS_pShc |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{102} = \text{vol}(c1) \cdot \text{J}110_{\text{k}} \cdot [\text{Grb2\_SOS\_pShc\_pTrkA\_endo}]$$
 (204)

Table 319: Properties of each parameter.

| Id     | Name | SBO | Value             | Unit | Constant |
|--------|------|-----|-------------------|------|----------|
| J110_k |      |     | $4.2\cdot10^{-4}$ |      |          |

### **5.103 Reaction** J112

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

Name pFRS2\_pTrkA\_endo\_degradation

### **Reaction equation**

$$pFRS2\_pTrkA\_endo \longrightarrow degradation + pFRS2$$
 (205)

### Reactant

Table 320: Properties of each reactant.

| Id               | Name | SBO |
|------------------|------|-----|
| pFRS2_pTrkA_endo |      |     |

## **Products**

Table 321: Properties of each product.

| Id                | Name | SBO |
|-------------------|------|-----|
| degradation pFRS2 |      |     |

## **Kinetic Law**

$$v_{103} = J112_k \cdot [pFRS2_pTrkA\_endo]$$
 (206)

Table 322: Properties of each parameter.

| Id     | Name | SBO | Value             | Unit | Constant |
|--------|------|-----|-------------------|------|----------|
| J112_k |      |     | $4.2\cdot10^{-4}$ |      |          |

### **5.104 Reaction** J113

This is a reversible reaction of one reactant forming one product.

Name form\_NGFreceptor

# **Reaction equation**

$$pro\_TrkA \Longrightarrow NGFR$$
 (207)

#### Reactant

Table 323: Properties of each reactant.

| Id       | Name | SBO |
|----------|------|-----|
| pro_TrkA |      |     |

### **Product**

Table 324: Properties of each product.

| Id   | Name | SBO |
|------|------|-----|
| NGFR |      |     |

#### **Kinetic Law**

$$v_{104} = vol(compartment) \cdot (J113_k1 \cdot [pro\_TrkA] - J113_k2 \cdot [NGFR])$$
 (208)

Table 325: Properties of each parameter.

| Id                 | Name | SBO | Value   | Unit | Constant |
|--------------------|------|-----|---|------|----------|
| J113_k1<br>J113_k2 |      | 2   | 8.333 · 10 <sup>-4</sup><br>2.7778 · 10 <sup>-4</sup> |      |          |

### **5.105 Reaction** J115

This is a reversible reaction of two reactants forming one product.

Name binding\_Shc\_to\_dpEGFR\_c\_Cbl

## **Reaction equation**

$$Shc + dpEGFR\_c\_Cbl \Longrightarrow Shc\_dpEGFR\_c\_Cbl$$
 (209)

### **Reactants**

Table 326: Properties of each reactant.

| Id                     | Name | SBO |
|------------------------|------|-----|
| Shc                    |      |     |
| ${\tt dpEGFR\_c\_Cbl}$ |      |     |

### **Product**

Table 327: Properties of each product.

| Id               | Name | SBO |
|------------------|------|-----|
| Shc_dpEGFR_c_Cbl |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{105} = \text{vol}(c1) \cdot (\text{J}115\text{\_k}1 \cdot [\text{Shc}] \cdot [\text{dpEGFR\_c\_Cbl}] - \text{J}115\text{\_k}2 \cdot [\text{Shc\_dpEGFR\_c\_Cbl}]) \quad (210)$$

Table 328: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant       |
|---------|------|----------------|----------------|
| J115_k1 |      | 10.0           | $\overline{Z}$ |
| J115_k2 |      | 0.2            | $\square$      |

### **5.106 Reaction** J116

This is a reversible reaction of two reactants forming one product.

Name binding\_pShc\_to\_dpEGFR\_c\_Cbl

### **Reaction equation**

$$dpEGFR\_c\_Cbl + pShc \Longrightarrow pShc\_dpEGFR\_c\_Cbl$$
 (211)

#### **Reactants**

Table 329: Properties of each reactant.

#### **Product**

Table 330: Properties of each product.

| Id                | Name | SBO |
|-------------------|------|-----|
| pShc_dpEGFR_c_Cbl |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{106} = \text{vol}\left(\text{c1}\right) \cdot \left(\text{J116\_k1} \cdot \left[\text{dpEGFR\_c\_Cbl}\right] \cdot \left[\text{pShc}\right] - \text{J116\_k2} \cdot \left[\text{pShc\_dpEGFR\_c\_Cbl}\right]\right) \quad (212)$$

Table 331: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant                |
|---------|------|----------------|-------------------------|
| J116_k1 |      | 10.0           | $\overline{\checkmark}$ |
| J116_k2 |      | 0.2            |                         |

### **5.107 Reaction** J117

This is a reversible reaction of two reactants forming one product.

Name binding\_SOS\_Grb2\_to\_pShc\_dpEGFR\_c\_Cbl

# **Reaction equation**

$$pShc\_dpEGFR\_c\_Cbl + SOS\_Grb2 \Longrightarrow Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl$$
 (213)

#### Reactants

Table 332: Properties of each reactant.

| Id Name SBO                   |  |  |
|-------------------------------|--|--|
| pShc_dpEGFR_c_Cbl<br>SOS_Grb2 |  |  |

#### **Product**

Table 333: Properties of each product.

|                            | 1    |     |
|----------------------------|------|-----|
| Id                         | Name | SBO |
| Grb2_SOS_pShc_dpEGFR_c_Cbl |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{107} = \text{vol}(c1) \cdot (\text{J}117_k1 \cdot [\text{pShc\_dpEGFR\_c\_Cbl}] \cdot [\text{SOS\_Grb2}] - \text{J}117_k2}$$

$$\cdot [\text{Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl}])$$
(214)

Table 334: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant  |
|---------|------|----------------|-----------|
| J117_k1 |      | 10.0           |           |
| J117_k2 |      | 0.2            | $\square$ |

### **5.108 Reaction** J119

This is a reversible reaction of two reactants forming one product.

Name binding\_c\_Cbl\_to\_Crk\_C3G\_pFRS2\_dpEGFR

## **Reaction equation**

$$c\_Cbl + Crk\_C3G\_pFRS2\_dpEGFR \Longrightarrow Crk\_C3G\_pFRS2\_dpEGFR\_c\_Cbl$$
 (215)

Table 335: Properties of each reactant.

| Tuest ever 1 repetities of euron 10 metunis. |      |     |  |
|--|------|-----|--|
| Id   | Name | SBO |  |
| c_Cbl<br>Crk_C3G_pFRS2_dpEGFR                |      |     |  |

Table 336: Properties of each product.

| Id                         | Name | SBO |
|----------------------------|------|-----|
| Crk_C3G_pFRS2_dpEGFR_c_Cbl |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{108} = vol(c1) \cdot (J119\_k1 \cdot [c\_Cbl] \cdot [Crk\_C3G\_pFRS2\_dpEGFR] - J119\_k2$$

$$\cdot [Crk\_C3G\_pFRS2\_dpEGFR\_c\_Cbl])$$

$$(216)$$

Table 337: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant                  |
|---------|------|----------------|---------------------------|
| J119_k1 |      | 0.5            | $ \overline{\checkmark} $ |
| J119_k2 |      | 0.2            |                           |

### **5.109 Reaction** J118

This is a reversible reaction of two reactants forming one product.

Name binding\_FRS2\_to\_dpEGFR\_c\_Cbl

# **Reaction equation**

$$dpEGFR_c\_Cbl + FRS2 \Longrightarrow FRS2\_dpEGFR_c\_Cbl$$
 (217)

Table 338: Properties of each reactant.

| Id                   | Name | SBO |
|----------------------|------|-----|
| dpEGFR_c_Cbl<br>FRS2 |      |     |

Table 339: Properties of each product.

| Id                | Name | SBO |
|-------------------|------|-----|
| FRS2_dpEGFR_c_Cbl |      |     |

#### **Kinetic Law**

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

$$v_{109} = \text{vol}(c1) \cdot (\text{J}118 \text{-k}1 \cdot [\text{dpEGFR}_c\text{-Cbl}] \cdot [\text{FRS2}] - \text{J}118 \text{-k}2 \cdot [\text{FRS2}\_\text{dpEGFR}_c\text{-Cbl}]) \quad (218)$$

Table 340: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant  |
|---------|------|----------------|-----------|
| J118_k1 |      | 1.0            |           |
| J118_k2 |      | 0.2            | $\square$ |

## **5.110 Reaction** J120

This is a reversible reaction of two reactants forming one product.

Name binding\_pFRS2\_to\_dpEGFR\_c\_Cbl

## **Reaction equation**

$$dpEGFR\_c\_Cbl + pFRS2 \Longrightarrow pFRS2\_dpEGFR\_c\_Cbl$$
 (219)

Table 341: Properties of each reactant.

| Id           | Name | SBO |
|--------------|------|-----|
| dpEGFR_c_Cbl |      |     |

| Id    | Name | SBO |
|-------|------|-----|
| pFRS2 |      |     |

Table 342: Properties of each product.

| Id                 | Name |  |
|--------------------|------|--|
| pFRS2_dpEGFR_c_Cb1 |      |  |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{110} = vol\left(c1\right) \cdot \left(J120\_k1 \cdot \left[dpEGFR\_c\_Cbl\right] \cdot \left[pFRS2\right] - J120\_k2 \cdot \left[pFRS2\_dpEGFR\_c\_Cbl\right]\right) \tag{220}$$

Table 343: Properties of each parameter.

|         |      | 1              |                |
|---------|------|----------------|----------------|
| Id      | Name | SBO Value Unit | Constant       |
| J120_k1 |      | 1.0            | $\overline{Z}$ |
| J120_k2 |      | 0.2            |                |

### **5.111 Reaction** J121

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

Name Ras\_GTP\_dephosphorylation

## **Reaction equation**

$$Ras\_GTP \xrightarrow{pDok\_RasGAP} Ras\_GDP$$
 (221)

Table 344: Properties of each reactant.

| Id      | Name | SBO |
|---------|------|-----|
| Ras_GTP |      |     |

#### Modifier

Table 345: Properties of each modifier.

| Id          | Name | SBO |
|-------------|------|-----|
| pDok_RasGAP |      |     |

#### **Product**

Table 346: Properties of each product.

| Id      | Name | SBO |
|---------|------|-----|
| Ras_GDP |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{111} = \text{vol}(c1) \cdot \frac{\text{J121\_Vmax} \cdot [\text{Ras\_GTP}] \cdot [\text{pDok\_RasGAP}]}{\text{J121\_Km1} + [\text{Ras\_GTP}]}$$
(222)

Table 347: Properties of each parameter.

| Id        | Name | SBO Value Unit | Constant       |
|-----------|------|----------------|----------------|
| J121_Vmax |      | 10.0           | $ \mathbf{Z} $ |
| J121_Km1  |      | 1.0            | $\square$      |

#### **5.112 Reaction** J122

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

Name RAP1\_GTP\_dephosphorylation

## **Reaction equation**

$$Rap1\_GTP \xrightarrow{Rap1GAP} Rap1\_GDP$$
 (223)

Table 348: Properties of each reactant.

| Id       | Name | SBO |
|----------|------|-----|
| Rap1_GTP |      |     |

### **Modifier**

Table 349: Properties of each modifier.

| Id      | Name | SBO |
|---------|------|-----|
| Rap1GAP |      |     |

### **Product**

Table 350: Properties of each product.

| Id       | Name | SBO |
|----------|------|-----|
| Rap1_GDP |      |     |

### **Kinetic Law**

Derived unit contains undeclared units

$$v_{112} = \text{vol}\left(\text{c1}\right) \cdot \frac{\text{J122\_Vmax} \cdot [\text{Rap1\_GTP}] \cdot [\text{Rap1GAP}]}{\text{J122\_Km1} + [\text{Rap1\_GTP}]} \tag{224}$$

Table 351: Properties of each parameter.

| Id        | Name | SBO | Value | Unit | Constant     |
|-----------|------|-----|-------|------|--------------|
| J122_Vmax |      |     | 2.0   |      | <b>✓</b>     |
| J122_Km1  |      |     | 1.0   |      | $\mathbf{Z}$ |

### **5.113 Reaction** J123

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

Name Dok\_phosphorylation

## **Reaction equation**

Dok pTrkA, Shc\_pTrkA, pShc\_pTrkA, Grb2\_SOS\_pShc\_pTrkA, FRS2\_pTrkA, pFRS2\_pTrkA, Crk\_C3G\_pFRS2\_pTrkA, (225)

#### Reactant

Table 352: Properties of each reactant.

| Id  | Name | SBO |
|-----|------|-----|
| Dok |      |     |

#### **Modifiers**

Table 353: Properties of each modifier.

| Id                  | Name | SBO |
|---------------------|------|-----|
| pTrkA               |      |     |
| Shc_pTrkA           |      |     |
| pShc_pTrkA          |      |     |
| Grb2_SOS_pShc_pTrkA |      |     |
| FRS2_pTrkA          |      |     |
| pFRS2_pTrkA         |      |     |
| Crk_C3G_pFRS2_pTrkA |      |     |

### **Product**

Table 354: Properties of each product.

| Id   | Name | SBO |
|------|------|-----|
| pDok |      |     |

### **Kinetic Law**

$$v_{113} = \text{vol} (c1) \tag{226} \\ \cdot \underbrace{J123\_V\text{max} \cdot [\text{Dok}] \cdot ([\text{pTrkA}] + [\text{Shc}\_\text{pTrkA}] + [\text{pShc}\_\text{pTrkA}] + [\text{Grb2}\_\text{SOS}\_\text{pShc}\_\text{pTrkA}] + [\text{FRS2}\_\text{pTrkA}] + [\text{Dok}]}$$

Table 355: Properties of each parameter.

| Id        | Name | SBO Value Unit | Constant       |
|-----------|------|----------------|----------------|
| J123_Vmax |      | 0.02           | $ \mathbf{Z} $ |
| J123_Km1  |      | 0.10           | $\checkmark$   |

#### **5.114 Reaction** J124

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

Name Grb1\_SOS\_pShc\_dissociation

# **Reaction equation**

Grb2\_SOS\_pShc 
$$\xrightarrow{\text{dppERK}}$$
 pShc + pSOS\_Grb2 (227)

#### Reactant

Table 356: Properties of each reactant.

| Id            | Name | SBO |
|---------------|------|-----|
| Grb2_SOS_pShc |      |     |

#### **Modifier**

Table 357: Properties of each modifier.

| Id     | Name | SBO |
|--------|------|-----|
| dppERK |      |     |

#### **Products**

Table 358: Properties of each product.

| Id                | Name | SBO |
|-------------------|------|-----|
| pShc<br>pSOS_Grb2 |      |     |

#### **Kinetic Law**

$$v_{114} = \text{vol}\left(\text{c1}\right) \cdot \frac{\text{J124\_Vmax} \cdot [\text{Grb2\_SOS\_pShc}] \cdot [\text{dppERK}]}{\text{J124\_Km1} + [\text{Grb2\_SOS\_pShc}]} \tag{228}$$

Table 359: Properties of each parameter.

| Id        | Name | SBO Value Unit | Constant     |
|-----------|------|----------------|--------------|
| J124_Vmax |      | 1.000          |              |
| J124_Km1  |      | 25.641         | $\checkmark$ |

### **5.115 Reaction** J133

This is a reversible reaction of two reactants forming one product.

Name binding\_MEK\_to\_ERK

### **Reaction equation**

$$ERK + MEK \Longrightarrow MEK\_ERK$$
 (229)

### **Reactants**

Table 360: Properties of each reactant.

| Id  | Name | SBO |
|-----|------|-----|
| ERK |      |     |
| MEK |      |     |

#### **Product**

Table 361: Properties of each product.

| Id      | Name | SBO |
|---------|------|-----|
| MEK_ERK |      |     |

#### **Kinetic Law**

$$v_{115} = \text{vol}(\text{c1}) \cdot (\text{J133\_k1} \cdot [\text{ERK}] \cdot [\text{MEK}] - \text{J133\_k2} \cdot [\text{MEK\_ERK}])$$
 (230)

Table 362: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant |
|---------|------|----------------|----------|
| J133_k1 |      | 16.304         |          |
| J133_k2 |      | 0.600          |          |

#### **5.116 Reaction** J134

This is a reversible reaction of two reactants forming one product.

Name binding\_ERK\_to\_pMEK

## **Reaction equation**

$$ERK + pMEK \Longrightarrow pMEK\_ERK$$
 (231)

#### **Reactants**

Table 363: Properties of each reactant.

| Id   | Name | SBO |
|------|------|-----|
| ERK  |      |     |
| рМЕК |      |     |

#### **Product**

Table 364: Properties of each product.

| Id       | Name | SBO |
|----------|------|-----|
| pMEK_ERK |      |     |

#### **Kinetic Law**

122

$$v_{116} = \text{vol}(\text{c1}) \cdot (\text{J134\_k1} \cdot [\text{ERK}] \cdot [\text{pMEK}] - \text{J134\_k2} \cdot [\text{pMEK\_ERK}])$$
 (232)

Table 365: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant       |
|---------|------|----------------|----------------|
| J134_k1 |      | 16.304         | $\overline{Z}$ |
| J134_k2 |      | 0.600          | $\checkmark$   |

### **5.117 Reaction** J135

This is a reversible reaction of two reactants forming one product.

Name binding\_ERK\_to\_ppMEK

## **Reaction equation**

$$ERK + ppMEK \Longrightarrow ppMEK\_ERK$$
 (233)

### **Reactants**

Table 366: Properties of each reactant.

| Id    | Name | SBO |
|-------|------|-----|
| ERK   |      |     |
| ppMEK |      |     |

### **Product**

Table 367: Properties of each product.

| Id        | Name | SBO |
|-----------|------|-----|
| ppMEK_ERK |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{117} = \text{vol}(\text{c1}) \cdot (\text{J135\_k1} \cdot [\text{ERK}] \cdot [\text{ppMEK}] - \text{J135\_k2} \cdot [\text{ppMEK\_ERK}])$$
 (234)

Table 368: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant |
|---------|------|----------------|----------|
| J135_k1 |      | 16.304         | Ø        |
| J135_k2 |      | 0.600          |          |

### **5.118 Reaction** J136

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

Name ppMEK\_ERK\_dissociation

### **Reaction equation**

$$ppMEK\_ERK \longrightarrow ppERK + ppMEK$$
 (235)

#### Reactant

Table 369: Properties of each reactant.

| Id        | Name | SBO |
|-----------|------|-----|
| ppMEK_ERK |      |     |

#### **Products**

Table 370: Properties of each product.

| Id             | Name | SBO |
|----------------|------|-----|
| ppERK<br>ppMEK |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{118} = \text{vol}(c1) \cdot \text{J}136\_\text{k} \cdot [\text{ppMEK\_ERK}]$$
 (236)

Table 371: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J136_k |      | 0.15           |          |

### **5.119 Reaction** J137

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

Name c\_Raf\_Ras\_GTP\_dissociation

## **Reaction equation**

$$c_Raf_Ras_GTP \xrightarrow{pDok_RasGAP} c_Raf_Ras_GDP$$
 (237)

Table 372: Properties of each reactant.

| Id            | Name | SBO |
|---------------|------|-----|
| c_Raf_Ras_GTP |      |     |

### **Modifier**

Table 373: Properties of each modifier.

| Id          | Name | SBO |
|-------------|------|-----|
| pDok_RasGAP |      |     |

### **Products**

Table 374: Properties of each product.

| Id                             | Name | SBO |
|--------------------------------|------|-----|
| c_Raf                          |      |     |
| $\mathtt{Ras}_{-}\mathtt{GDP}$ |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{119} = \text{vol}(c1) \cdot \frac{\text{J137\_Vmax} \cdot [\text{c\_Raf\_Ras\_GTP}] \cdot [\text{pDok\_RasGAP}]}{\text{J137\_Km1} + [\text{c\_Raf\_Ras\_GTP}]}$$
(238)

Table 375: Properties of each parameter.

| Id        | Name | SBO Value Unit | Constant  |
|-----------|------|----------------|-----------|
| J137_Vmax |      | 10.0           |           |
| J137_Km1  |      | 1.0            | $\square$ |

## **5.120 Reaction** J138

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

Name B\_Raf\_Ras\_GTP\_dissociation

## **Reaction equation**

$$B\_Raf\_Ras\_GTP \xrightarrow{pDok\_RasGAP} B\_Raf + Ras\_GDP$$
 (239)

### Reactant

Table 376: Properties of each reactant.

| Id            | Name | SBO |
|---------------|------|-----|
| B_Raf_Ras_GTP |      |     |

### **Modifier**

Table 377: Properties of each modifier.

| Id          | Name | SBO |
|-------------|------|-----|
| pDok_RasGAP |      |     |

### **Products**

Table 378: Properties of each product.

| Id         | Name | SBO |
|------------|------|-----|
| B_Raf      |      |     |
| $Ras\_GDP$ |      |     |

#### **Kinetic Law**

$$v_{120} = vol(c1) \cdot \frac{J138\_Vmax \cdot [B\_Raf\_Ras\_GTP] \cdot [pDok\_RasGAP]}{J138\_Km1 + [B\_Raf\_Ras\_GTP]} \tag{240}$$

Table 379: Properties of each parameter.

| Id        | Name | SBO Value Unit | Constant     |
|-----------|------|----------------|--------------|
| J138_Vmax |      | 10.0           |              |
| J138_Km1  |      | 1.0            | $\checkmark$ |

### **5.121 Reaction** J139

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

Name B\_Raf\_Rap1\_GTP\_dissociation

## **Reaction equation**

$$B\_Raf\_Rap1\_GTP \xrightarrow{Rap1GAP} B\_Raf + Rap1\_GDP$$
 (241)

#### Reactant

Table 380: Properties of each reactant.

| Id             | Name | SBO |
|----------------|------|-----|
| B_Raf_Rap1_GTP |      |     |

### **Modifier**

Table 381: Properties of each modifier.

| Id      | Name | SBO |
|---------|------|-----|
| Rap1GAP |      |     |

### **Products**

Table 382: Properties of each product.

| Id          | Name | SBO |
|-------------|------|-----|
| B_Raf       |      |     |
| $Rap1\_GDP$ |      |     |

### **Kinetic Law**

$$v_{121} = vol(c1) \cdot \frac{J139\_Vmax \cdot [B\_Raf\_Rap1\_GTP] \cdot [Rap1GAP]}{J139\_Km1 + [B\_Raf\_Rap1\_GTP]}$$
(242)

Table 383: Properties of each parameter.

| Id        | Name | SBO Value Unit | Constant     |
|-----------|------|----------------|--------------|
| J139_Vmax |      | 2.0            |              |
| J139_Km1  |      | 1.0            | $\checkmark$ |

### **5.122 Reaction** J140

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

## **Reaction equation**

$$c_Raf_Ras_GTP + MEK \longrightarrow c_Raf_Ras_GTP_MEK$$
 (243)

#### **Reactants**

Table 384: Properties of each reactant.

| Id            | Name | SBO |
|---------------|------|-----|
| c_Raf_Ras_GTP |      |     |
| MEK           |      |     |

### **Product**

Table 385: Properties of each product.

| Id                | Name | SBO |
|-------------------|------|-----|
| c_Raf_Ras_GTP_MEK |      |     |

### **Kinetic Law**

$$v_{122} = \text{vol}(c1) \cdot (\text{J}140 \text{\_k}1 \cdot [\text{c\_Raf\_Ras\_GTP}] \cdot [\text{MEK}] - \text{J}140 \text{\_k}2 \cdot [\text{c\_Raf\_Ras\_GTP\_MEK}]) \quad (244)$$

Table 386: Properties of each parameter.

| Id      | Name | SBO | Value  | Unit | Constant        |
|---------|------|-----|--------|------|-----------------|
| J140_k1 |      |     | 15.625 |      | $ \mathcal{A} $ |
| J140_k2 |      |     | 2.000  |      |                 |

### **5.123 Reaction** J141

This is a reversible reaction of two reactants forming one product.

## **Reaction equation**

$$c_Raf_Ras_GTP + pMEK \rightleftharpoons c_Raf_Ras_GTP_pMEK$$
 (245)

#### **Reactants**

Table 387: Properties of each reactant.

| Id                    | Name | SBO |
|-----------------------|------|-----|
| c_Raf_Ras_GTP<br>pMEK |      |     |

#### **Product**

Table 388: Properties of each product.

| Id                 | Name | SBO |
|--------------------|------|-----|
| c_Raf_Ras_GTP_pMEK |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{123} = vol\left(c1\right) \cdot \left(J141\_k1 \cdot \left[c\_Raf\_Ras\_GTP\right] \cdot \left[pMEK\right] - J141\_k2 \cdot \left[c\_Raf\_Ras\_GTP\_pMEK\right]\right) \tag{246}$$

Table 389: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant       |
|---------|------|----------------|----------------|
| J141_k1 |      | 15.625         | $\overline{Z}$ |
| J141_k2 |      | 2.000          | $ \mathbf{Z} $ |

### **5.124 Reaction** J142

This is a reversible reaction of two reactants forming one product.

## **Reaction equation**

$$c\_Raf\_Ras\_GTP + MEK\_ERK \Longrightarrow c\_Raf\_Ras\_GTP\_MEK\_ERK$$
 (247)

### **Reactants**

Table 390: Properties of each reactant.

| Id            | Name | SBO |
|---------------|------|-----|
| c_Raf_Ras_GTP |      |     |
| MEK_ERK       |      |     |

#### **Product**

Table 391: Properties of each product.

| Id                    | Name | SBO |
|-----------------------|------|-----|
| c_Raf_Ras_GTP_MEK_ERK |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{124} = \text{vol}(\text{c1}) \cdot (\text{J142\_k1} \cdot [\text{c\_Raf\_Ras\_GTP}] \cdot [\text{MEK\_ERK}] - \text{J142\_k2}$$

$$\cdot [\text{c\_Raf\_Ras\_GTP\_MEK\_ERK}])$$
(248)

Table 392: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant        |
|---------|------|----------------|-----------------|
| J142_k1 |      | 15.625         | $ \mathcal{A} $ |
| J142_k2 |      | 2.000          | $\checkmark$    |

### **5.125 Reaction** J143

This is a reversible reaction of two reactants forming one product.

## **Reaction equation**

$$c_Raf_Ras_GTP + pMEK_ERK \Longrightarrow c_Raf_Ras_GTP_pMEK_ERK$$
 (249)

#### **Reactants**

Table 393: Properties of each reactant.

| Id                        | Name | SBO |
|---------------------------|------|-----|
| c_Raf_Ras_GTP<br>pMEK_ERK |      |     |

#### **Product**

Table 394: Properties of each product.

| Id                     | Name | SBO |
|------------------------|------|-----|
| c_Raf_Ras_GTP_pMEK_ERK |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{125} = vol(c1) \cdot (J143\_k1 \cdot [c\_Raf\_Ras\_GTP] \cdot [pMEK\_ERK] - J143\_k2$$

$$\cdot [c\_Raf\_Ras\_GTP\_pMEK\_ERK])$$
(250)

Table 395: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant       |
|---------|------|----------------|----------------|
| J143_k1 |      | 15.625         | $ \mathbf{Z} $ |
| J143_k2 |      | 2.000          | $\square$      |

### **5.126 Reaction** J144

This is a reversible reaction of two reactants forming one product.

## **Reaction equation**

$$B_Raf_Ras_GTP + MEK \Longrightarrow B_Raf_Ras_GTP_MEK$$
 (251)

Table 396: Properties of each reactant.

| Id            | Name | SBO |
|---------------|------|-----|
| B_Raf_Ras_GTP |      |     |
| MEK           |      |     |

Table 397: Properties of each product.

| Id                | Name | SBO |
|-------------------|------|-----|
| B_Raf_Ras_GTP_MEK |      |     |

#### **Kinetic Law**

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

$$v_{126} = vol\left(c1\right)\cdot\left(J144\_k1\cdot\left[B\_Raf\_Ras\_GTP\right]\cdot\left[MEK\right] - J144\_k2\cdot\left[B\_Raf\_Ras\_GTP\_MEK\right]\right) \tag{252}$$

Table 398: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant       |
|---------|------|----------------|----------------|
| J144_k1 |      | 6.25           | $\overline{Z}$ |
| J144_k2 |      | 0.80           | $\square$      |

## **5.127 Reaction** J145

This is a reversible reaction of two reactants forming one product.

## **Reaction equation**

$$B_Raf_Ras_GTP + pMEK \Longrightarrow B_Raf_Ras_GTP_pMEK$$
 (253)

Table 399: Properties of each reactant.

| Id            | Name | SBO |
|---------------|------|-----|
| B_Raf_Ras_GTP |      |     |
| pMEK          |      |     |

| Id | Name | SBO |
|----|------|-----|
|    |      |     |

Table 400: Properties of each product.

| Id                 | Name | SBO |
|--------------------|------|-----|
| B_Raf_Ras_GTP_pMEK |      |     |

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{127} = vol\left(c1\right) \cdot \left(J145\_k1 \cdot \left[B\_Raf\_Ras\_GTP\right] \cdot \left[pMEK\right] - J145\_k2 \cdot \left[B\_Raf\_Ras\_GTP\_pMEK\right]\right) \tag{254}$$

Table 401: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant |
|---------|------|----------------|----------|
| J145_k1 |      | 6.25           |          |
| J145_k2 |      | 0.80           |          |

### **5.128 Reaction** J146

This is a reversible reaction of two reactants forming one product.

## **Reaction equation**

$$B\_Raf\_Ras\_GTP + MEK\_ERK \Longrightarrow B\_Raf\_Ras\_GTP\_MEK\_ERK$$
 (255)

#### **Reactants**

Table 402: Properties of each reactant.

| Id                       | Name | SBO |
|--------------------------|------|-----|
| B_Raf_Ras_GTP<br>MEK_ERK |      |     |

## **Product**

Table 403: Properties of each product.

| Id                    | Name |  |
|-----------------------|------|--|
| B_Raf_Ras_GTP_MEK_ERK |      |  |

### **Kinetic Law**

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

$$v_{128} = vol(c1) \cdot (J146\_k1 \cdot [B\_Raf\_Ras\_GTP] \cdot [MEK\_ERK] - J146\_k2$$

$$\cdot [B\_Raf\_Ras\_GTP\_MEK\_ERK])$$
(256)

Table 404: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant       |
|---------|------|----------------|----------------|
| J146_k1 |      | 6.25           | $\overline{Z}$ |
| J146_k2 |      | 0.80           | $\checkmark$   |

### **5.129 Reaction** J147

This is a reversible reaction of two reactants forming one product.

## **Reaction equation**

$$B\_Raf\_Ras\_GTP + pMEK\_ERK \Longrightarrow B\_Raf\_Ras\_GTP\_pMEK\_ERK$$
 (257)

## **Reactants**

Table 405: Properties of each reactant.

| Id            | Name | SBO |
|---------------|------|-----|
| B_Raf_Ras_GTP |      |     |
| pMEK_ERK      |      |     |

### **Product**

Table 406: Properties of each product.

| Id                     | Name | SBO |
|------------------------|------|-----|
| B_Raf_Ras_GTP_pMEK_ERK |      |     |

### **Kinetic Law**

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

$$v_{129} = vol(c1) \cdot (J147\_k1 \cdot [B\_Raf\_Ras\_GTP] \cdot [pMEK\_ERK] - J147\_k2$$

$$\cdot [B\_Raf\_Ras\_GTP\_pMEK\_ERK])$$
(258)

Table 407: Properties of each parameter.

| Id        | Name | SBO Value Unit | Constant   |
|-----------|------|----------------|------------|
| J147_k1   |      | 6.25           | lacksquare |
| $J147_k2$ |      | 0.80           | $\square$  |

### **5.130 Reaction** J148

This is a reversible reaction of two reactants forming one product.

## **Reaction equation**

$$B_Raf_Rap1_GTP + MEK \Longrightarrow B_Raf_Rap1_GTP\_MEK$$
 (259)

#### **Reactants**

Table 408: Properties of each reactant.

| Id             | Name | SBO |
|----------------|------|-----|
| B_Raf_Rap1_GTP |      |     |
| MEK            |      |     |

#### **Product**

Table 409: Properties of each product.

| Id                 | Name | SBO |
|--------------------|------|-----|
| B_Raf_Rap1_GTP_MEK |      |     |

## **Kinetic Law**

$$v_{130} = vol\left(c1\right) \cdot \left(J148\_k1 \cdot \left[B\_Raf\_Rap1\_GTP\right] \cdot \left[MEK\right] - J148\_k2 \cdot \left[B\_Raf\_Rap1\_GTP\_MEK\right]\right) \tag{260}$$

Table 410: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant                     |
|---------|------|----------------|------------------------------|
| J148_k1 |      | 9.375          |                              |
| J148_k2 |      | 1.200          | $   \overline{\mathscr{L}} $ |

### **5.131 Reaction** J149

This is a reversible reaction of two reactants forming one product.

## **Reaction equation**

$$B_Raf_Rap1_GTP + pMEK \Longrightarrow B_Raf_Rap1_GTP_pMEK$$
 (261)

#### **Reactants**

Table 411: Properties of each reactant.

| Id             | Name | SBO |
|----------------|------|-----|
| B_Raf_Rap1_GTP |      |     |
| рМЕК           |      |     |

### **Product**

Table 412: Properties of each product.

| 1                   | 1    |     |
|---------------------|------|-----|
| Id                  | Name | SBO |
| B_Raf_Rap1_GTP_pMEK |      |     |

### **Kinetic Law**

$$v_{131} = vol\left(c1\right) \cdot \left(J149\_k1 \cdot \left[B\_Raf\_Rap1\_GTP\right] \cdot \left[pMEK\right] - J149\_k2 \cdot \left[B\_Raf\_Rap1\_GTP\_pMEK\right]\right) \tag{262}$$

Table 413: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant     |
|---------|------|----------------|--------------|
| J149_k1 |      | 9.375          |              |
| J149_k2 |      | 1.200          | $\checkmark$ |

#### **5.132 Reaction** J150

This is a reversible reaction of two reactants forming one product.

## **Reaction equation**

$$B_Raf_Rap1_GTP + MEK_ERK \Longrightarrow B_Raf_Rap1_GTP_MEK_ERK$$
 (263)

#### **Reactants**

Table 414: Properties of each reactant.

| Id             | Name | SBO |
|----------------|------|-----|
| B_Raf_Rap1_GTP |      |     |
| MEK_ERK        |      |     |

### **Product**

Table 415: Properties of each product.

| Id                     | Name | SBO |
|------------------------|------|-----|
| B_Raf_Rap1_GTP_MEK_ERK |      |     |

### **Kinetic Law**

$$v_{132} = \text{vol}(\text{c1}) \cdot (\text{J150\_k1} \cdot [\text{B\_Raf\_Rap1\_GTP}] \cdot [\text{MEK\_ERK}] - \text{J150\_k2}$$

$$\cdot [\text{B\_Raf\_Rap1\_GTP\_MEK\_ERK}])$$
(264)

Table 416: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant  |
|---------|------|----------------|-----------|
| J150_k1 |      | 9.375          |           |
| J150_k2 |      | 1.200          | $\square$ |

### **5.133 Reaction** J151

This is a reversible reaction of two reactants forming one product.

## **Reaction equation**

$$B_Raf_Rap1_GTP + pMEK_ERK \Longrightarrow B_Raf_Rap1_GTP_pMEK_ERK$$
 (265)

#### **Reactants**

Table 417: Properties of each reactant.

| Id                         | Name | SBO |
|----------------------------|------|-----|
| B_Raf_Rap1_GTP<br>pMEK_ERK |      |     |

#### **Product**

Table 418: Properties of each product.

| Id                      | Name | SBO |
|-------------------------|------|-----|
| B_Raf_Rap1_GTP_pMEK_ERK |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$\begin{aligned} \nu_{133} &= vol\left(c1\right) \cdot \left(J151\_k1 \cdot \left[B\_Raf\_Rap1\_GTP\right] \cdot \left[pMEK\_ERK\right] - J151\_k2 \right. \\ & \cdot \left[B\_Raf\_Rap1\_GTP\_pMEK\_ERK\right] \right) \end{aligned} \tag{266}$$

Table 419: Properties of each parameter.

| Id                 | Name | SBO Value Unit | Constant |
|--------------------|------|----------------|----------|
| J151_k1<br>J151_k2 |      | 9.375<br>1.200 | <b>✓</b> |
| J151_k2            |      | 1.200          |          |

#### **5.134 Reaction** J152

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

### **Reaction equation**

$$c_Raf_Ras_GTP_MEK \longrightarrow c_Raf_Ras_GTP + pMEK$$
 (267)

#### Reactant

Table 420: Properties of each reactant.

| THE IZE IZE ITE PETERS | ,, ,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,, |     |
|------------------------|---|-----|
| Id                     | Name                                    | SBO |
| c_Raf_Ras_GTP_MEK      |   |     |

### **Products**

Table 421: Properties of each product.

| Id            | Name | SBO |
|---------------|------|-----|
| c_Raf_Ras_GTP |      |     |
| pMEK          |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{134} = \text{vol}(c1) \cdot \text{J}152\_k \cdot [c\_Raf\_Ras\_GTP\_MEK]$$
 (268)

Table 422: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J152_k |      | 0.5            |          |

## **5.135 Reaction** J153

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

## **Reaction equation**

$$c\_Raf\_Ras\_GTP\_pMEK \longrightarrow c\_Raf\_Ras\_GTP + ppMEK$$
 (269)

| Table 423: Properties of each reactant. |
|---|
|---|

| Tuble 423. Troperties of each reactant. |      |     |  |  |
|---|------|-----|--|--|
| Id                                      | Name | SBO |  |  |
| c_Raf_Ras_GTP_pMEK                      |      |     |  |  |

Table 424: Properties of each product.

| Id            | Name | SBO |
|---------------|------|-----|
| c_Raf_Ras_GTP |      |     |
| ррМЕК         |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{135} = \text{vol}(c1) \cdot \text{J}153\_\text{k} \cdot [c\_\text{Raf}\_\text{Ras}\_\text{GTP}\_\text{pMEK}]$$
 (270)

Table 425: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J153_k |      | 0.5            |          |

## **5.136 Reaction** J154

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

## **Reaction equation**

$$c_Raf_Ras_GTP\_MEK\_ERK \longrightarrow c_Raf_Ras_GTP + pMEK\_ERK$$
 (271)

### Reactant

Table 426: Properties of each reactant.

| Id                    | Name | SBO |
|-----------------------|------|-----|
| c_Raf_Ras_GTP_MEK_ERK |      |     |

#### **Products**

Table 427: Properties of each product.

| Id                        | Name | SBO |
|---------------------------|------|-----|
| c_Raf_Ras_GTP<br>pMEK_ERK |      |     |

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{136} = \text{vol}(c1) \cdot \text{J}154\_\text{k} \cdot [c\_\text{Raf}\_\text{Ras}\_\text{GTP}\_\text{MEK}\_\text{ERK}]$$
 (272)

Table 428: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J154_k |      | 0.5            |          |

### **5.137 Reaction** J155

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

# **Reaction equation**

$$c_Raf_Ras_GTP_pMEK_ERK \longrightarrow c_Raf_Ras_GTP + ppMEK_ERK$$
 (273)

#### Reactant

Table 429: Properties of each reactant.

| Tuote (2) (Troperties of e |      |     |
|----------------------------|------|-----|
| Id                         | Name | SBO |
| c_Raf_Ras_GTP_pMEK_ERK     |      |     |

### **Products**

Table 430: Properties of each product.

| Id            | Name | SBO |
|---------------|------|-----|
| c_Raf_Ras_GTP |      |     |
| ppMEK_ERK     |      |     |

### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{137} = \text{vol}(c1) \cdot \text{J}155_k \cdot [c_Raf_Ras_GTP_pMEK\_ERK]$$
 (274)

Table 431: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J155_k |      | 0.5            | Ø        |

## **5.138 Reaction** J156

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

### **Reaction equation**

$$B\_Raf\_Ras\_GTP\_MEK \longrightarrow B\_Raf\_Ras\_GTP + pMEK$$
 (275)

#### Reactant

Table 432: Properties of each reactant.

| Id                | Name | SBO |
|-------------------|------|-----|
| B_Raf_Ras_GTP_MEK |      |     |

### **Products**

Table 433: Properties of each product.

| B_Raf_Ras_GTP<br>pMEK |  |
|-----------------------|--|

### **Kinetic Law**

$$v_{138} = \text{vol}(c1) \cdot \text{J}156\_\text{k} \cdot [\text{B\_Raf\_Ras\_GTP\_MEK}]$$
 (276)

Table 434: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J156_k |      | 0.2            |          |

### **5.139 Reaction** J157

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

# **Reaction equation**

$$B\_Raf\_Ras\_GTP\_pMEK \longrightarrow B\_Raf\_Ras\_GTP + ppMEK$$
 (277)

#### Reactant

Table 435: Properties of each reactant.

| Id                 | Name | SBO |
|--------------------|------|-----|
| B_Raf_Ras_GTP_pMEK |      |     |

### **Products**

Table 436: Properties of each product.

| Id            | Name | SBO |
|---------------|------|-----|
| B_Raf_Ras_GTP |      |     |
| ppMEK         |      |     |

### **Kinetic Law**

$$v_{139} = \text{vol}(c1) \cdot \text{J}157\_\text{k} \cdot [B\_\text{Raf}\_\text{Ras}\_\text{GTP}\_\text{pMEK}]$$
 (278)

Table 437: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J157_k |      | 0.2            |          |

## **5.140 Reaction** J158

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

# **Reaction equation**

$$B_Raf_Ras_GTP_MEK_ERK \longrightarrow B_Raf_Ras_GTP + pMEK_ERK$$
 (279)

#### Reactant

Table 438: Properties of each reactant.

| Id                    | Name | SBO |
|-----------------------|------|-----|
| B_Raf_Ras_GTP_MEK_ERK |      |     |

#### **Products**

Table 439: Properties of each product.

| Id                        | Name | SBO |
|---------------------------|------|-----|
| B_Raf_Ras_GTP<br>pMEK_ERK |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{140} = \text{vol}(c1) \cdot \text{J}158\text{-k} \cdot [\text{B\_Raf\_Ras\_GTP\_MEK\_ERK}]$$
 (280)

Table 440: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant     |
|--------|------|----------------|--------------|
| J158_k |      | 0.2            | $\checkmark$ |

#### **5.141 Reaction** J159

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

# **Reaction equation**

$$B_Raf_Ras_GTP_pMEK_ERK \longrightarrow B_Raf_Ras_GTP + ppMEK_ERK$$
 (281)

#### Reactant

Table 441: Properties of each reactant.

| Table 441. I Toperties of Cach reactant. |      |     |
|--|------|-----|
| Id                                       | Name | SBO |
| B_Raf_Ras_GTP_pMEK_ERK                   |      |     |

## **Products**

Table 442: Properties of each product.

| Id            | Name | SBO |
|---------------|------|-----|
| B_Raf_Ras_GTP |      |     |
| ppMEK_ERK     |      |     |

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{141} = \text{vol}(c1) \cdot \text{J}159 \text{-k} \cdot [\text{B\_Raf\_Ras\_GTP\_pMEK\_ERK}]$$
 (282)

Table 443: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J159_k |      | 0.2            |          |

# **5.142 Reaction** J160

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

# **Reaction equation**

$$B_Raf_Rap1_GTP_MEK \longrightarrow B_Raf_Rap1_GTP + pMEK$$
 (283)

# Reactant

Table 444: Properties of each reactant.

| Id                 | Name | SBO |
|--------------------|------|-----|
| B_Raf_Rap1_GTP_MEK |      |     |

## **Products**

Table 445: Properties of each product.

| Id                  | Name | SBO |
|---------------------|------|-----|
| B_Raf_Rap1_GTP pMEK |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{142} = \text{vol}(c1) \cdot \text{J}160\text{\_k} \cdot [\text{B\_Raf\_Rap1\_GTP\_MEK}]$$
 (284)

Table 446: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J160_k |      | 0.3            |          |

# **5.143 Reaction** J161

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

# **Reaction equation**

$$B\_Raf\_Rap1\_GTP\_pMEK \longrightarrow B\_Raf\_Rap1\_GTP + ppMEK$$
 (285)

## Reactant

Table 447: Properties of each reactant.

| Id                  | Name | SBO |
|---------------------|------|-----|
| B_Raf_Rap1_GTP_pMEK |      |     |

# **Products**

Table 448: Properties of each product.

| Id             | Name | SBO |
|----------------|------|-----|
| B_Raf_Rap1_GTP |      |     |

| Id    | Name | SBO |
|-------|------|-----|
| ррМЕК |      |     |

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{143} = \text{vol}(c1) \cdot \text{J}161\_\text{k} \cdot [\text{B\_Raf\_Rap1\_GTP\_pMEK}]$$
 (286)

Table 449: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J161_k |      | 0.3            |          |

# **5.144 Reaction** J162

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

# **Reaction equation**

$$B\_Raf\_Rap1\_GTP\_MEK\_ERK \longrightarrow B\_Raf\_Rap1\_GTP + pMEK\_ERK \qquad (287)$$

## Reactant

Table 450: Properties of each reactant.

| Id                     | Name |  |
|------------------------|------|--|
| B_Raf_Rap1_GTP_MEK_ERK |      |  |

## **Products**

Table 451: Properties of each product.

| Id             | Name | SBO |
|----------------|------|-----|
| B_Raf_Rap1_GTP |      |     |
| pMEK_ERK       |      |     |

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{144} = \text{vol}(c1) \cdot \text{J}162 \cdot \text{k} \cdot [\text{B\_Raf\_Rap1\_GTP\_MEK\_ERK}]$$
 (288)

Table 452: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J162_k |      | 0.3            |          |

#### **5.145 Reaction** J163

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

# **Reaction equation**

$$B_Raf_Rap1_GTP_pMEK_ERK \longrightarrow B_Raf_Rap1_GTP + ppMEK_ERK$$
 (289)

#### Reactant

Table 453: Properties of each reactant.

| Id                      | Name | SBO |
|-------------------------|------|-----|
| B_Raf_Rap1_GTP_pMEK_ERK |      |     |

#### **Products**

Table 454: Properties of each product.

| Name | SBO  |
|------|------|
|      |      |
|      | Name |

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{145} = \text{vol}(c1) \cdot \text{J}163 \cdot \text{k} \cdot [\text{B\_Raf\_Rap1\_GTP\_pMEK\_ERK}]$$
 (290)

Table 455: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J163_k |      | 0.3            |          |

## **5.146 Reaction** J164

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

# **Reaction equation**

$$Crk\_C3G\_pFRS2\_dpEGFR\_c\_Cbl\_ubiq \longrightarrow c\_Cbl + pFRS2 + Crk\_C3G \tag{291}$$

#### Reactant

Table 456: Properties of each reactant.

| Id                              | Name | SBO |
|---------------------------------|------|-----|
| Crk_C3G_pFRS2_dpEGFR_c_Cbl_ubiq |      |     |

## **Products**

Table 457: Properties of each product.

| Id               | Name | SBO |
|------------------|------|-----|
| c_Cbl<br>pFRS2   |      |     |
| ${\tt Crk\_C3G}$ |      |     |

#### **Kinetic Law**

Derived unit contains undeclared units

$$v_{146} = vol(c1) \cdot J164\_k \cdot [Crk\_C3G\_pFRS2\_dpEGFR\_c\_Cbl\_ubiq]$$
 (292)

Table 458: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant     |
|--------|------|----------------|--------------|
| J164_k |      | 0.001          | $\checkmark$ |

# **5.147 Reaction** J165

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

# **Reaction equation**

$$MKP3 + dppERK \longrightarrow dppERK\_MKP3$$
 (293)

#### **Reactants**

Table 459: Properties of each reactant.

| Id     | Name | SBO |
|--------|------|-----|
| МКРЗ   |      |     |
| dppERK |      |     |

#### **Product**

Table 460: Properties of each product.

| Id          | Name | SBO |
|-------------|------|-----|
| dppERK_MKP3 |      |     |

## **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{147} = \text{vol}\left(\text{c1}\right) \cdot \left(\text{J165\_k1} \cdot \left[\text{MKP3}\right] \cdot \left[\text{dppERK}\right] - \text{J165\_k2} \cdot \left[\text{dppERK\_MKP3}\right]\right) \quad (294)$$

Table 461: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant |
|---------|------|----------------|----------|
| J165_k1 |      | 15.00          |          |
| J165_k2 |      | 0.24           |          |

## **5.148 Reaction** J166

This is a reversible reaction of two reactants forming one product.

# **Reaction equation**

$$MKP3 + ppERK \Longrightarrow ppERK\_MKP3$$
 (295)

#### **Reactants**

Table 462: Properties of each reactant.

| Id    | Name | SBO |
|-------|------|-----|
| МКРЗ  |      |     |
| ppERK |      |     |

## **Product**

Table 463: Properties of each product.

| Id         | Name | SBO |
|------------|------|-----|
| ppERK_MKP3 |      |     |

# **Kinetic Law**

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

$$v_{148} = vol(c1) \cdot (J166 k1 \cdot [MKP3] \cdot [ppERK] - J166 k2 \cdot [ppERK MKP3])$$
 (296)

Table 464: Properties of each parameter.

| Id      | Name | SBO Value Unit | Constant     |
|---------|------|----------------|--------------|
| J166_k1 |      | 15.00          |              |
| J166_k2 |      | 0.24           | $\checkmark$ |

# **5.149 Reaction** J167

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

# **Reaction equation**

$$ppERK\_MKP3 \longrightarrow ERK + MKP3 \tag{297}$$

#### Reactant

Table 465: Properties of each reactant.

| Id         | Name | SBO |
|------------|------|-----|
| ppERK_MKP3 |      |     |

## **Products**

Table 466: Properties of each product.

| Id   | Name | SBO |
|------|------|-----|
| ERK  |      |     |
| MKP3 |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{149} = \text{vol}(c1) \cdot \text{J}167_{-}\text{k} \cdot [\text{ppERK\_MKP3}]$$
 (298)

Table 467: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J167_k |      | 0.06           |          |

## **5.150 Reaction** J168

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

# **Reaction equation**

$$dppERK\_MKP3 \longrightarrow ppERK + ERK + MKP3 \tag{299}$$

## Reactant

Table 468: Properties of each reactant.

| Id          | Name | SBO |
|-------------|------|-----|
| dppERK_MKP3 |      |     |

# **Products**

Table 469: Properties of each product.

| Id    | Name | SBO |
|-------|------|-----|
| ppERK |      |     |

| Id          | Name | SBO |
|-------------|------|-----|
| ERK<br>MKP3 |      |     |

#### **Kinetic Law**

**Derived unit** contains undeclared units

$$v_{150} = \text{vol}(c1) \cdot \text{J}168 \cdot \text{k} \cdot [\text{dppERK\_MKP3}] \tag{300}$$

Table 470: Properties of each parameter.

| Id     | Name | SBO Value Unit | Constant |
|--------|------|----------------|----------|
| J168_k |      | 0.06           |          |

# **6 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.

## **6.1 Species** EGFR

Initial concentration  $0.3 \, \mu \text{mol} \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in re2 and as a product in re1).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{EGFR} = |v_1| - |v_2| \tag{301}$$

# 6.2 Species L\_EGFR

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

This species takes part in two reactions (as a reactant in re8 and as a product in re2).

$$\frac{\mathrm{d}}{\mathrm{d}t} L \cdot \text{EGFR} = |v_2| - 2|v_3| \tag{302}$$

## 6.3 Species L\_EGFR\_dimer

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

This species takes part in two reactions (as a reactant in J5 and as a product in re8).

$$\frac{d}{dt}L\_EGFR\_dimer = |v_3| - |v_6|$$
 (303)

## **6.4 Species SOS**

Initial concentration  $0.1 \, \mu mol \cdot l^{-1}$ 

This species takes part in three reactions (as a reactant in J3, J51 and as a product in J10).

$$\frac{d}{dt}SOS = |v_{11}| - |v_4| - |v_{50}| \tag{304}$$

## 6.5 Species L\_dpEGFR

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

This species takes part in eight reactions (as a reactant in J6, J7, J12, J23, J35, J36 and as a product in J5 and as a modifier in J31).

$$\frac{d}{dt}L_{dp}EGFR = |v_{6}| - |v_{7}| - |v_{8}| - |v_{13}| - |v_{24}| - |v_{35}| - |v_{36}|$$
(305)

# 6.6 Species pSOS

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

This species takes part in three reactions (as a reactant in J4, J10 and as a product in J51).

$$\frac{d}{dt}pSOS = |v_{50}| - |v_{5}| - |v_{11}| \tag{306}$$

# 6.7 Species SOS\_Grb2

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

This species takes part in nine reactions (as a reactant in J22, J24, J50, J95, J96, J117 and as a product in J3, J11, J30).

$$\frac{d}{dt}SOS\_Grb2 = v_4 + |v_{12}| + |v_{30}| - |v_{23}| - |v_{25}| - |v_{49}| - |v_{87}| - |v_{88}| - |v_{107}|$$
(307)

# 6.8 Species Grb2

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

This species takes part in two reactions (as a reactant in J3, J4).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Grb2} = -|v_4| - |v_5| \tag{308}$$

# 6.9 Species Dok

Initial concentration  $0.3 \ \mu mol \cdot l^{-1}$ 

This species takes part in three reactions (as a reactant in J31, J123 and as a product in J8).

$$\frac{d}{dt}Dok = |v_9| - |v_{31}| - |v_{113}| \tag{309}$$

# 6.10 Species pDok

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

This species takes part in four reactions (as a reactant in J8, J49 and as a product in J31, J123).

$$\frac{d}{dt}pDok = |v_{31}| + |v_{113}| - v_9 - |v_{48}|$$
(310)

## 6.11 Species Crk

Initial concentration 1 µmol·1<sup>-1</sup>

This species takes part in one reaction (as a reactant in J34).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Crk} = -v_{34} \tag{311}$$

## **6.12 Species** FRS2

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

This species takes part in eight reactions (as a reactant in J35, J77, J84, J118 and as a product in J33, J46, J87, J92).

$$\frac{d}{dt}FRS2 = |v_{33}| + |v_{46}| + |v_{80}| + |v_{84}| - |v_{35}| - |v_{70}| - |v_{77}| - |v_{109}|$$
(312)

# 6.13 Species Shc

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

This species takes part in nine reactions (as a reactant in J12, J75, J79, J115 and as a product in J18, J30, J32, J89, J93).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Shc} = |v_{19}| + |v_{30}| + |v_{32}| + |v_{82}| + |v_{85}| - |v_{13}| - |v_{68}| - |v_{72}| - |v_{105}| \tag{313}$$

# 6.14 Species pSOS\_Grb2

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

This species takes part in four reactions (as a reactant in J11 and as a product in J4, J50, J124).

$$\frac{d}{dt}pSOS\_Grb2 = |v_5| + |v_{49}| + |v_{114}| - |v_{12}|$$
(314)

#### 6.15 Species Rap1\_GDP

Initial concentration  $0.2 \ \mu mol \cdot l^{-1}$ 

This species takes part in four reactions (as a reactant in J68 and as a product in J67, J122, J139).

$$\frac{d}{dt}Rap1\_GDP = |v_{60}| + |v_{112}| + |v_{121}| - |v_{61}|$$
(315)

#### 6.16 Species MEK

Initial concentration  $0.68 \ \mu mol \cdot l^{-1}$ 

This species takes part in five reactions (as a reactant in J133, J140, J144, J148 and as a product in J58).

$$\frac{d}{dt}MEK = v_{55} - |v_{115}| - |v_{122}| - |v_{126}| - |v_{130}|$$
(316)

## 6.17 Species MKP3

Initial concentration  $0.018 \ \mu mol \cdot l^{-1}$ 

This species takes part in four reactions (as a reactant in J165, J166 and as a product in J167, J168).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{MKP3} = |v_{149}| + |v_{150}| - |v_{147}| - |v_{148}| \tag{317}$$

## 6.18 Species pShc\_dpEGFR

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

This species takes part in five reactions (as a reactant in J9, J24 and as a product in J7, J13 and as a modifier in J31).

$$\frac{d}{dt}pShc\_dpEGFR = |v_8| + |v_{14}| - |v_{10}| - |v_{25}|$$
(318)

## 6.19 Species dpEGFR\_c\_Cbl

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

This species takes part in eight reactions (as a reactant in J14, J27, J115, J116, J118, J120 and as a product in J6 and as a modifier in J31).

$$\frac{d}{dt}dpEGFR_cC_Cbl = |v_7| - |v_{15}| - |v_{27}| - |v_{105}| - |v_{106}| - |v_{109}| - |v_{110}|$$
(319)

# 6.20 Species B\_Raf\_Rap1\_GTP

# Initial concentration $0 \, \mu mol \cdot l^{-1}$

This species takes part in ten reactions (as a reactant in J139, J148, J149, J150, J151 and as a product in J53, J160, J161, J162, J163).

$$\frac{d}{dt}B_Raf_Rap1_GTP = v_{52} + v_{142} + v_{143} + v_{144} + v_{145} - v_{121} - v_{130} - v_{131} - v_{132} - v_{133}$$
(320)

#### 6.21 Species pShc\_dpEGFR\_c\_Cbl

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

This species takes part in six reactions (as a reactant in J19, J117 and as a product in J9, J21, J116 and as a modifier in J31).

$$\frac{d}{dt}pShc_dpEGFR_c_CCbl = |v_{10}| + |v_{22}| + |v_{106}| - |v_{20}| - |v_{107}|$$
(321)

# **6.22 Species** pFRS2\_dpEGFR\_c\_Cbl

#### Initial concentration $0 \, \mu mol \cdot l^{-1}$

This species takes part in five reactions (as a reactant in J41, J44 and as a product in J40, J43, J120).

$$\frac{d}{dt} pFRS2\_dpEGFR\_c\_Cbl = |v_{40}| + |v_{43}| + |v_{110}| - |v_{41}| - |v_{44}|$$
(322)

## 6.23 Species Shc\_dpEGFR

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

This species takes part in four reactions (as a reactant in J13, J16 and as a product in J12 and as a modifier in J31).

$$\frac{d}{dt}Shc\_dpEGFR = v_{13} - |v_{14}| - |v_{17}|$$
 (323)

#### 6.24 Species c\_Cbl

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

This species takes part in 14 reactions (as a reactant in J6, J9, J16, J25, J39, J40, J119 and as a product in J15, J18, J20, J29, J46, J47, J164).

$$\frac{d}{dt}c\_Cbl = v_{16} + v_{19} + v_{21} + v_{29} + v_{46} + v_{47} + v_{146} - v_7 - v_{10} - v_{17} - v_{26} - v_{39} - v_{40} - v_{108}$$
(324)

#### 6.25 Species RasGAP

# Initial concentration $0.1~\mu mol \cdot l^{-1}$

This species takes part in one reaction (as a reactant in J49).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{RasGAP} = -v_{48} \tag{325}$$

## **6.26 Species** c\_Raf

# Initial concentration $0.5 \, \mu mol \cdot l^{-1}$

This species takes part in two reactions (as a reactant in J52 and as a product in J137).

$$\frac{d}{dt}c Raf = |v_{119}| - |v_{51}| \tag{326}$$

# **6.27 Species** B\_Raf

# Initial concentration $0.2~\mu mol \cdot l^{-1}$

This species takes part in four reactions (as a reactant in J53, J54 and as a product in J138, J139).

$$\frac{\mathrm{d}}{\mathrm{d}t} \mathbf{B} \cdot \mathbf{Raf} = |v_{120}| + |v_{121}| - |v_{52}| - |v_{53}| \tag{327}$$

## 6.28 Species ERK

Initial concentration  $0.26 \, \mu \text{mol} \cdot l^{-1}$ 

This species takes part in five reactions (as a reactant in J133, J134, J135 and as a product in J167, J168).

$$\frac{\mathrm{d}}{\mathrm{d}t} \text{ERK} = |v_{149}| + |v_{150}| - |v_{115}| - |v_{116}| - |v_{117}| \tag{328}$$

# 6.29 Species PP2A

Initial concentration  $0.24 \ \mu mol \cdot l^{-1}$ 

This species takes part in four reactions (as a modifier in J57, J58, J61, J62).

$$\frac{\mathrm{d}}{\mathrm{d}t}PP2A = 0 \tag{329}$$

#### 6.30 Species Ras\_GDP

Initial concentration  $0.1 \, \mu mol \cdot l^{-1}$ 

This species takes part in five reactions (as a reactant in J69 and as a product in J66, J121, J137, J138).

$$\frac{d}{dt}Ras\_GDP = |v_{59}| + |v_{111}| + |v_{119}| + |v_{120}| - |v_{62}|$$
(330)

# 6.31 Species Rap1GAP

Initial concentration  $0.012 \ \mu mol \cdot l^{-1}$ 

This species takes part in two reactions (as a modifier in J122, J139).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Rap}1\mathrm{GAP} = 0\tag{331}$$

# 6.32 Species C3G

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

This species takes part in one reaction (as a reactant in J34).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{C3G} = -v_{34} \tag{332}$$

# 6.33 Species NGFR

Initial concentration  $0.061894 \ \mu mol \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in J70 and as a product in J113).

$$\frac{\mathrm{d}}{\mathrm{d}t} \mathrm{NGFR} = |v_{104}| - |v_{63}| \tag{333}$$

#### 6.34 Species pShc

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

This species takes part in ten reactions (as a reactant in J7, J22, J32, J76, J80, J116 and as a product in J20, J90, J94, J124).

$$\frac{d}{dt}pShc = |v_{21}| + |v_{83}| + |v_{86}| + |v_{114}| - |v_{8}| - |v_{23}| - |v_{32}| - |v_{69}| - |v_{73}| - |v_{106}|$$
(334)

## 6.35 Species pFRS2\_dpEGFR

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

This species takes part in five reactions (as a reactant in J38, J40 and as a product in J36, J37 and as a modifier in J31).

$$\frac{d}{dt} pFRS2\_dpEGFR = |v_{36}| + |v_{37}| - |v_{38}| - |v_{40}|$$
(335)

# 6.36 Species pTrkA\_endo

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

This species takes part in seven reactions (as a reactant in J73, J79, J80, J84, J85, J106 and as a product in J72).

$$\frac{d}{dt} p Trk A\_endo = |v_{65}| - |v_{66}| - |v_{72}| - |v_{73}| - |v_{77}| - |v_{78}| - |v_{98}|$$
(336)

## 6.37 Species MEK\_ERK

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

This species takes part in five reactions (as a reactant in J142, J146, J150 and as a product in J62, J133).

$$\frac{d}{dt}MEK\_ERK = |v_{57}| + |v_{115}| - |v_{124}| - |v_{128}| - |v_{132}|$$
(337)

## 6.38 Species pMEK\_ERK

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

This species takes part in nine reactions (as a reactant in J62, J143, J147, J151 and as a product in J61, J134, J154, J158, J162).

$$\frac{d}{dt}pMEK\_ERK = |v_{56}| + |v_{116}| + |v_{136}| + |v_{140}| + |v_{144}| - |v_{57}| - |v_{125}| - |v_{129}| - |v_{133}|$$
(338)

# 6.39 Species FRS2\_dpEGFR\_c\_Cbl\_ubiq

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

This species takes part in two reactions (as a reactant in J46 and as a product in J42).

$$\frac{d}{dt}FRS2\_dpEGFR\_c\_Cbl\_ubiq = v_{42} - v_{46}$$
(339)

# 6.40 Species Crk\_C3G\_pFRS2\_dpEGFR\_c\_Cbl

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

This species takes part in five reactions (as a reactant in J45 and as a product in J44, J119 and as a modifier in J31, J68).

$$\frac{d}{dt} Crk_{C3G_pFRS2_dpEGFR_cC_Cbl} = |v_{44}| + |v_{108}| - |v_{45}|$$
(340)

## **6.41 Species** pShc\_dpEGFR\_c\_Cbl\_ubiq

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

This species takes part in two reactions (as a reactant in J20 and as a product in J19).

$$\frac{d}{dt}pShc\_dpEGFR\_c\_Cbl\_ubiq = v_{20} - v_{21}$$
(341)

## 6.42 Species Crk\_C3G\_pFRS2\_dpEGFR

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

This species takes part in four reactions (as a reactant in J119 and as a product in J38 and as a modifier in J31, J68).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Crk}_{-}\mathrm{C3G}_{-}\mathrm{pFRS2}_{-}\mathrm{dpEGFR} = v_{38} - v_{108}$$
 (342)

## **6.43 Species** Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl\_ubiq

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

This species takes part in two reactions (as a reactant in J29 and as a product in J28).

$$\frac{d}{dt}Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl\_ubiq = v_{28} - v_{29}$$
(343)

# **6.44 Species** Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl

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

This species takes part in six reactions (as a reactant in J28 and as a product in J25, J27, J117 and as a modifier in J31, J69).

$$\frac{d}{dt}Grb2\_SOS\_pShc\_dpEGFR\_c\_Cbl = |v_{26}| + |v_{27}| + |v_{107}| - |v_{28}|$$
(344)

# 6.45 Species Shc\_dpEGFR\_c\_Cbl\_ubiq

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

This species takes part in two reactions (as a reactant in J18 and as a product in J17).

$$\frac{d}{dt}Shc\_dpEGFR\_c\_Cbl\_ubiq = v_{18} - v_{19}$$
(345)

#### **6.46 Species** dpEGFR\_c\_Cbl\_ubiq

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

This species takes part in two reactions (as a reactant in J15 and as a product in J14).

$$\frac{d}{dt}dpEGFR_c\_Cbl\_ubiq = |v_{15}| - |v_{16}|$$
(346)

#### **6.47 Species** proteosome

Name proteasome

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

This species takes part in six reactions (as a product in J15, J18, J20, J29, J46, J47).

$$\frac{d}{dt} \text{proteosome} = v_{16} + v_{19} + v_{21} + v_{29} + v_{46} + v_{47}$$
(347)

# 6.48 Species Grb2\_SOS\_pShc

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

This species takes part in ten reactions (as a reactant in J23, J27, J30, J105, J106, J124 and as a product in J22, J29, J109, J110).

$$\frac{d}{dt}Grb2\_SOS\_pShc = v_{23} + v_{29} + v_{101} + v_{102} - v_{24} - v_{27} - v_{30} - v_{97} - v_{98} - v_{114}$$
(348)

## 6.49 Species Shc\_dpEGFR\_c\_Cbl

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

This species takes part in five reactions (as a reactant in J17, J21 and as a product in J16, J115 and as a modifier in J31).

$$\frac{d}{dt}Shc_dpEGFR_cC_Cbl = |v_{17}| + |v_{105}| - |v_{18}| - |v_{22}|$$
(349)

# **6.50** Species Grb2\_SOS\_pShc\_dpEGFR

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

This species takes part in five reactions (as a reactant in J25 and as a product in J23, J24 and as a modifier in J31, J69).

$$\frac{d}{dt}Grb2\_SOS\_pShc\_dpEGFR = |v_{24}| + |v_{25}| - |v_{26}|$$
(350)

## **6.51 Species** pFRS2

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

This species takes part in eleven reactions (as a reactant in J33, J36, J78, J85, J120 and as a product in J47, J88, J107, J108, J112, J164).

$$\frac{d}{dt}pFRS2 = v_{47} + v_{81} + v_{99} + v_{100} + v_{103} + v_{146} - v_{33} - v_{36} - v_{71} - v_{78} - v_{110}$$
(351)

# **6.52** Species FRS2\_dpEGFR

#### Initial concentration $0 \, \mu mol \cdot l^{-1}$

This species takes part in four reactions (as a reactant in J37, J39 and as a product in J35 and as a modifier in J31).

$$\frac{d}{dt}FRS2\_dpEGFR = |v_{35}| - |v_{37}| - |v_{39}|$$
(352)

# 6.53 Species pDok\_RasGAP

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

This species takes part in four reactions (as a product in J49 and as a modifier in J121, J137, J138).

$$\frac{d}{dt}pDok\_RasGAP = v_{48}$$
 (353)

# 6.54 Species pMEK

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

This species takes part in nine reactions (as a reactant in J58, J134, J141, J145, J149 and as a product in J57, J152, J156, J160).

$$\frac{d}{dt}pMEK = v_{54} + v_{134} + v_{138} + v_{142} - v_{55} - v_{116} - v_{123} - v_{127} - v_{131}$$
 (354)

## 6.55 Species FRS2\_dpEGFR\_c\_Cb1

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

This species takes part in five reactions (as a reactant in J42, J43 and as a product in J39, J118 and as a modifier in J31).

$$\frac{d}{dt}FRS2\_dpEGFR\_c\_Cbl = |v_{39}| + |v_{109}| - |v_{42}| - |v_{43}|$$
(355)

## 6.56 Species pFRS2\_dpEGFR\_c\_Cbl\_ubiq

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

This species takes part in two reactions (as a reactant in J47 and as a product in J41).

$$\frac{d}{dt} pFRS2\_dpEGFR\_c\_Cbl\_ubiq = v_{41} - v_{47}$$
(356)

#### 6.57 Species Ras\_GTP

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

This species takes part in five reactions (as a reactant in J52, J54, J66, J121 and as a product in J69).

$$\frac{d}{dt} Ras\_GTP = |v_{62}| - |v_{51}| - |v_{53}| - |v_{59}| - |v_{111}|$$
(357)

# 6.58 Species Crk\_C3G\_pFRS2\_dpEGFR\_c\_Cbl\_ubiq

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

This species takes part in two reactions (as a reactant in J164 and as a product in J45).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{Crk}_{C3}\mathrm{G}_{p}\mathrm{FRS2}_{d}\mathrm{p}\mathrm{EGFR}_{c}_{C}\mathrm{Cbl}_{u}\mathrm{biq} = v_{45} - v_{146}$$
(358)

# 6.59 Species c\_Raf\_Ras\_GTP

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

This species takes part in ten reactions (as a reactant in J137, J140, J141, J142, J143 and as a product in J52, J152, J153, J154, J155).

$$\frac{d}{dt}c_Raf_Ras_GTP = v_{51} + v_{134} + v_{135} + v_{136} + v_{137} - v_{119} - v_{122} - v_{123} - v_{124} - v_{125}$$
(359)

#### 6.60 Species B\_Raf\_Ras\_GTP

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

This species takes part in ten reactions (as a reactant in J138, J144, J145, J146, J147 and as a product in J54, J156, J157, J158, J159).

$$\frac{d}{dt}B_Raf_Ras_GTP = v_{53} + v_{138} + v_{139} + v_{140} + v_{141} - v_{120} - v_{120} - v_{126} - v_{127} - v_{128} - v_{129}$$
(360)

## **6.61 Species** ppMEK

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

This species takes part in six reactions (as a reactant in J57, J135 and as a product in J136, J153, J157, J161).

$$\frac{d}{dt}ppMEK = |v_{118}| + |v_{135}| + |v_{139}| + |v_{143}| - |v_{54}| - |v_{117}|$$
(361)

## 6.62 Species ppERK

#### Initial concentration $0 \, \mu mol \cdot l^{-1}$

This species takes part in four reactions (as a reactant in J63, J166 and as a product in J136, J168).

$$\frac{\mathrm{d}}{\mathrm{d}t} \mathrm{ppERK} = |v_{118}| + |v_{150}| - 2|v_{58}| - |v_{148}| \tag{362}$$

# 6.63 Species pTrkA

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

This species takes part in nine reactions (as a reactant in J72, J74, J75, J76, J77, J78, J105 and as a product in J71 and as a modifier in J123).

$$\frac{d}{dt}pTrkA = v_{64} - v_{65} - v_{67} - v_{68} - v_{69} - v_{70} - v_{71} - v_{97}$$
(363)

## 6.64 Species Crk\_C3G

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

This species takes part in eight reactions (as a reactant in J38, J44, J103, J104 and as a product in J34, J107, J108, J164).

$$\frac{d}{dt} \text{Crk\_C3G} = |v_{34}| + |v_{99}| + |v_{100}| + |v_{146}| - |v_{38}| - |v_{44}| - |v_{95}| - |v_{96}|$$
(364)

#### 6.65 Species Rap1\_GTP

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

This species takes part in four reactions (as a reactant in J53, J67, J122 and as a product in J68).

$$\frac{d}{dt}Rap1\_GTP = |v_{61}| - |v_{52}| - |v_{60}| - |v_{112}|$$
(365)

## 6.66 Species L\_NGFR

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

This species takes part in two reactions (as a reactant in J71 and as a product in J70).

$$\frac{\mathrm{d}}{\mathrm{d}t} L_{\mathrm{NGFR}} = v_{63} - v_{64} \tag{366}$$

## **6.67 Species** ppMEK\_ERK

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

This species takes part in six reactions (as a reactant in J61, J136 and as a product in J135, J155, J159, J163).

$$\frac{d}{dt}ppMEK\_ERK = |v_{117}| + |v_{137}| + |v_{141}| + |v_{145}| - |v_{56}| - |v_{118}|$$
(367)

# 6.68 Species dppERK

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

This species takes part in five reactions (as a reactant in J165 and as a product in J63 and as a modifier in J50, J51, J124).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{dppERK} = |v_{58}| - |v_{147}| \tag{368}$$

# 6.69 Species Shc\_pTrkA

#### Initial concentration $0 \, \mu mol \cdot l^{-1}$

This species takes part in five reactions (as a reactant in J82, J89, J102 and as a product in J75 and as a modifier in J123).

$$\frac{d}{dt}Shc_pTrkA = v_{68} - |v_{75}| - |v_{82}| - |v_{94}|$$
(369)

## 6.70 Species Shc\_pTrkA\_endo

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

This species takes part in four reactions (as a reactant in J81, J93 and as a product in J79, J102).

$$\frac{d}{dt} Shc_p Trk A_e ndo = v_{72} + v_{94} - v_{74} - v_{85}$$
 (370)

## 6.71 Species pShc\_pTrkA

#### Initial concentration $0 \, \mu mol \cdot l^{-1}$

This species takes part in six reactions (as a reactant in J90, J96, J101 and as a product in J76, J82 and as a modifier in J123).

$$\frac{d}{dt}pShc_pTrkA = |v_{69}| + |v_{75}| - |v_{83}| - |v_{88}| - |v_{93}|$$
(371)

# 6.72 Species pFRS2\_pTrkA

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

This species takes part in six reactions (as a reactant in J88, J99, J103 and as a product in J78, J83 and as a modifier in J123).

$$\frac{d}{dt}pFRS2\_pTrkA = |v_{71}| + |v_{76}| - |v_{81}| - |v_{91}| - |v_{95}|$$
(372)

## 6.73 Species FRS2\_pTrkA

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

This species takes part in five reactions (as a reactant in J83, J87, J100 and as a product in J77 and as a modifier in J123).

$$\frac{d}{dt}FRS2_pTrkA = |v_{70}| - |v_{76}| - |v_{80}| - |v_{92}|$$
(373)

# 6.74 Species pShc\_pTrkA\_endo

#### Initial concentration $0 \, \mu mol \cdot l^{-1}$

This species takes part in five reactions (as a reactant in J94, J95 and as a product in J80, J81, J101).

$$\frac{d}{dt}pShc_pTrkA_endo = v_{73} + v_{74} + v_{93} - v_{86} - v_{87}$$
 (374)

# 6.75 Species FRS2\_pTrkA\_endo

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

This species takes part in four reactions (as a reactant in J86, J92 and as a product in J84, J100).

$$\frac{d}{dt}FRS2_pTrkA\_endo = v_{77} + v_{92} - v_{79} - v_{84}$$
 (375)

## 6.76 Species pFRS2\_pTrkA\_endo

#### Initial concentration $0 \, \mu mol \cdot l^{-1}$

This species takes part in five reactions (as a reactant in J104, J112 and as a product in J85, J86, J99).

$$\frac{d}{dt} pFRS2_p TrkA_e ndo = |v_{78}| + |v_{79}| + |v_{91}| - |v_{96}| - |v_{103}|$$
(376)

# 6.77 Species Crk\_C3G\_pFRS2\_pTrkA\_endo

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

This species takes part in four reactions (as a reactant in J108 and as a product in J98, J104 and as a modifier in J68).

$$\frac{d}{dt} Crk_{-}C3G_{-}pFRS2_{-}pTrkA_{-}endo = |v_{90}| + |v_{96}| - |v_{100}|$$
(377)

# 6.78 Species Grb2\_SOS\_pShc\_pTrkA

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

This species takes part in six reactions (as a reactant in J97, J109 and as a product in J96, J105 and as a modifier in J69, J123).

$$\frac{d}{dt}Grb2\_SOS\_pShc\_pTrkA = |v_{88}| + |v_{97}| - |v_{89}| - |v_{101}|$$
(378)

# 6.79 Species Crk\_C3G\_pFRS2\_pTrkA

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

This species takes part in four reactions (as a reactant in J98, J107 and as a product in J103 and as a modifier in J123).

$$\frac{d}{dt} Crk_{-}C3G_{-}pFRS2_{-}pTrkA = v_{95} - v_{90} - v_{99}$$
(379)

# 6.80 Species Grb2\_SOS\_pShc\_pTrkA\_endo

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

This species takes part in four reactions (as a reactant in J110 and as a product in J95, J97, J106).

$$\frac{d}{dt}Grb2\_SOS\_pShc\_pTrkA\_endo = |v_{87}| + |v_{89}| + |v_{98}| - |v_{102}|$$
(380)

# 6.81 Species c\_Raf\_Ras\_GTP\_MEK

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

This species takes part in two reactions (as a reactant in J152 and as a product in J140).

$$\frac{\mathrm{d}}{\mathrm{d}t} c_{\mathrm{Raf}_{\mathrm{Ras}_{\mathrm{GTP}_{\mathrm{MEK}}}} = v_{122} - v_{134}$$
 (381)

## 6.82 Species c\_Raf\_Ras\_GTP\_pMEK

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

This species takes part in two reactions (as a reactant in J153 and as a product in J141).

$$\frac{\mathrm{d}}{\mathrm{d}t} c_{\mathrm{R}} - \mathrm{Ras} - \mathrm{GTP} - \mathrm{pMEK} = |v_{123}| - |v_{135}|$$
(382)

#### 6.83 Species c\_Raf\_Ras\_GTP\_MEK\_ERK

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

This species takes part in two reactions (as a reactant in J154 and as a product in J142).

$$\frac{\mathrm{d}}{\mathrm{d}t} c_{\mathrm{Raf}_{\mathrm{Ras}_{\mathrm{G}}}} \operatorname{GTP\_MEK\_ERK} = |v_{124}| - |v_{136}|$$
(383)

## 6.84 Species c\_Raf\_Ras\_GTP\_pMEK\_ERK

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

This species takes part in two reactions (as a reactant in J155 and as a product in J143).

$$\frac{\mathrm{d}}{\mathrm{d}t} c_{\mathrm{R}} - \mathrm{Ras}_{\mathrm{G}} - \mathrm{Ras}_{\mathrm{G}} - \mathrm{PMEK}_{\mathrm{E}} - \mathrm{RK} = v_{125} - v_{137}$$
(384)

# **6.85 Species** B\_Raf\_Ras\_GTP\_MEK

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

This species takes part in two reactions (as a reactant in J156 and as a product in J144).

$$\frac{\mathrm{d}}{\mathrm{d}t} \mathbf{B}_{\mathbf{R}} \mathbf{as}_{\mathbf{G}} \mathbf{TP}_{\mathbf{M}} \mathbf{E} \mathbf{K} = |v_{126}| - |v_{138}| \tag{385}$$

## 6.86 Species B\_Raf\_Ras\_GTP\_pMEK

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

This species takes part in two reactions (as a reactant in J157 and as a product in J145).

$$\frac{\mathrm{d}}{\mathrm{d}t} \mathbf{B}_{\mathbf{R}} \mathbf{as}_{\mathbf{G}} \mathbf{TP}_{\mathbf{p}} \mathbf{MEK} = v_{127} - v_{139}$$
(386)

## 6.87 Species B\_Raf\_Ras\_GTP\_MEK\_ERK

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

This species takes part in two reactions (as a reactant in J158 and as a product in J146).

$$\frac{d}{dt}B_Raf_Ras_GTP_MEK_ERK = v_{128} - v_{140}$$
 (387)

# 6.88 Species B\_Raf\_Ras\_GTP\_pMEK\_ERK

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

This species takes part in two reactions (as a reactant in J159 and as a product in J147).

$$\frac{\mathrm{d}}{\mathrm{d}t} \mathbf{B} \cdot \mathbf{Raf} \cdot \mathbf{Ras} \cdot \mathbf{GTP} \cdot \mathbf{pMEK} \cdot \mathbf{ERK} = |v_{129}| - |v_{141}|$$
(388)

# 6.89 Species B\_Raf\_Rap1\_GTP\_MEK

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

This species takes part in two reactions (as a reactant in J160 and as a product in J148).

$$\frac{d}{dt}B_{Raf}Rap1_{GTP}MEK = v_{130} - v_{142}$$
 (389)

## 6.90 Species B\_Raf\_Rap1\_GTP\_pMEK

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

This species takes part in two reactions (as a reactant in J161 and as a product in J149).

$$\frac{d}{dt} B_R af_R ap1_G TP_p MEK = v_{131} - v_{143}$$
 (390)

# **6.91 Species** B\_Raf\_Rap1\_GTP\_MEK\_ERK

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

This species takes part in two reactions (as a reactant in J162 and as a product in J150).

$$\frac{\mathrm{d}}{\mathrm{d}t} \mathbf{B}_{\mathbf{R}} \mathbf{a} \mathbf{f}_{\mathbf{R}} \mathbf{a} \mathbf{p} \mathbf{1}_{\mathbf{G}} \mathbf{T} \mathbf{P}_{\mathbf{M}} \mathbf{E} \mathbf{K}_{\mathbf{E}} \mathbf{R} \mathbf{K} = v_{132} - v_{144}$$
(391)

## 6.92 Species B\_Raf\_Rap1\_GTP\_pMEK\_ERK

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

This species takes part in two reactions (as a reactant in J163 and as a product in J151).

$$\frac{\mathrm{d}}{\mathrm{d}t} \mathbf{B}_{\mathbf{R}} \mathbf{a} \mathbf{f}_{\mathbf{R}} \mathbf{a} \mathbf{p} \mathbf{1}_{\mathbf{G}} \mathbf{T} \mathbf{P}_{\mathbf{p}} \mathbf{M} \mathbf{E} \mathbf{K}_{\mathbf{E}} \mathbf{R} \mathbf{K} = |\mathbf{v}_{133}| - |\mathbf{v}_{145}|$$
(392)

## 6.93 Species ppERK\_MKP3

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

This species takes part in two reactions (as a reactant in J167 and as a product in J166).

$$\frac{d}{dt} ppERK_MKP3 = |v_{148}| - |v_{149}|$$
(393)

# **6.94 Species** dppERK\_MKP3

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

This species takes part in two reactions (as a reactant in J168 and as a product in J165).

$$\frac{d}{dt}dppERK\_MKP3 = v_{147} - v_{150}$$
 (394)

#### 6.95 Species pro\_TrkA

Initial concentration  $0.020631 \, \mu \text{mol} \cdot l^{-1}$ 

This species takes part in one reaction (as a reactant in J113), which does not influence its rate of change because this species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{pro}_{-}\mathrm{TrkA} = 0 \tag{395}$$

## 6.96 Species NGF

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

This species takes part in one reaction (as a reactant in J70), which does not influence its rate of change because this species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{NGF} = 0\tag{396}$$

## 6.97 Species EGF

Initial concentration  $0.001613 \ \mu mol \cdot l^{-1}$ 

This species takes part in one reaction (as a reactant in re2), which does not influence its rate of change because this species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{EGF} = 0\tag{397}$$

# 6.98 Species pro\_EGFR

Initial concentration  $0.3~\mu mol \cdot l^{-1}$ 

This species takes part in one reaction (as a reactant in re1), which does not influence its rate of change because this species is on the boundary of the reaction system:

$$\frac{d}{dt} \text{pro\_EGFR} = 0 \tag{398}$$

## 6.99 Species degradation

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

This species takes part in 14 reactions (as a product in J73, J74, J87, J88, J89, J90, J92, J93, J94, J107, J108, J109, J110, J112), which do not influence its rate of change because this species is on the boundary of the reaction system:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{degradation} = 0 \tag{399}$$

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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