# **SBML Model Report**

# Model name: "Kirschner1998\_Immunotherapy\_Tumour"



May 6, 2016

# 1 General Overview

This is a document in SBML Level 2 Version 4 format. This model was created by the following three authors: Catherine Lloyd<sup>1</sup>, Catherine Lloyd<sup>2</sup> and Catherine Lloyd<sup>3</sup> at June 25<sup>th</sup> 2010 at 12:43 a.m. and last time modified at June 25<sup>th</sup> 2010 at 12:43 a.m. Table 1 gives an overview of the quantities of all components of this model.

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

| Element           | Quantity | Element              | Quantity |
|-------------------|----------|----------------------|----------|
| compartment types | 0        | compartments         | 1        |
| species types     | 0        | species              | 0        |
| events            | 0        | constraints          | 0        |
| reactions         | 0        | function definitions | 0        |
| global parameters | 17       | unit definitions     | 5        |
| rules             | 3        | initial assignments  | 0        |

#### **Model Notes**

This a model from the article:

Modeling immunotherapy of the tumor-immune interaction.

Kirschner D, Panetta JC. J Math Biol 1998 Sep;37(3):235-52 9785481,

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#### **Abstract:**

A number of lines of evidence suggest that immunotherapy with the cytokineinterleukin-2 (IL-2) may boost the immune system to fight tumors. CD4+ T cells, the cells that orchestrate the immune response, use these cytokines as signalingmechanisms for immune-response stimulation as well as lymphocyte stimulation, growth, and differentiation. Because tumor cells begin as 'self', the immunesystem may not respond in an effective way to eradicate them. Adoptive cellularimmunotherapy can potentially restore or enhance these effects. We illustratethrough mathematical modeling the dynamics between tumor cells, immune-effectorcells, and IL-2. These efforts are able to explain both short tumor oscillations in tumor sizes as well as long-term tumor relapse. We then explore the effectsof adoptive cellular immunotherapy on the model and describe under whatcircumstances the tumor can be eliminated.

This model was taken from the CellML repository and automatically converted to SBML. The original model was: **Kirschner D, Panetta JC.** (1998) - version=1.0 The original CellML model was created by:

#### **Catherine Lloyd**

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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 ten unit definitions of which five are predefined by SBML and not mentioned in the model.

#### 2.1 Unit day

Name day

**Definition** 86400 s

#### 2.2 Unit first\_order\_rate\_constant

Name first\_order\_rate\_constant

**Definition**  $(86400 \text{ s})^{-1}$ 

# 2.3 Unit cells\_per\_microlitre

Name cells\_per\_microlitre

**Definition**  $\mu l^{-1}$ 

#### 2.4 Unit flux

Name flux

**Definition**  $\mu l^{-1} \cdot (86400 \text{ s})^{-1}$ 

#### 2.5 Unit micromolar

Name micromolar

**Definition**  $\mu mol \cdot l^{-1}$ 

#### 2.6 Unit substance

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

**Definition** mol

#### 2.7 Unit volume

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

**Definition** 1

## 2.8 Unit area

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

**Definition**  $m^2$ 

# 2.9 Unit length

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

**Definition** m

## 2.10 Unit time

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

**Definition** s

# 3 Compartment

This model contains one compartment.

Table 2: Properties of all compartments.

| Id          | Name | SBO | Spatial Dimensions | Size | Unit | Constant | Outside |
|-------------|------|-----|--------------------|------|------|----------|---------|
| COMpartment |      |     | 3                  | 1    |      |          |         |

# 3.1 Compartment COMpartment

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

# 4 Parameters

This model contains 17 global parameters.

Table 3: Properties of each parameter.

| Id  | Name | SBO | Value             | Unit          | Constant       |
|-----|------|-----|-------------------|---------------|----------------|
| tau | tau  |     | 0.000             | dimensionless | $\overline{Z}$ |
| mu2 | mu2  |     | 0.030             | dimensionless |                |
| С   | c    |     | 0.025             | dimensionless |                |
| p1  | p1   |     | 0.125             | dimensionless |                |
| g1  | g1   |     | $2 \cdot 10^{-7}$ | dimensionless |                |
| s1  | s1   |     | 0.000             | dimensionless |                |
| x   | X    |     | 0.000             | dimensionless |                |
| У   | У    |     | 1.000             | dimensionless |                |
| r2  | r2   |     | 0.180             | dimensionless |                |
| a   | a    |     | 1.000             | dimensionless |                |
| b   | b    |     | $10^{-9}$         | dimensionless |                |
| g2  | g2   |     | 100000.000        | dimensionless | $\square$      |
| z   | Z    |     | 0.000             | dimensionless |                |
| mu3 | mu3  |     | 10.000            | dimensionless | $\square$      |
| p2  | p2   |     | 5.000             | dimensionless | $\square$      |
| g3  | g3   |     | 1000.000          | dimensionless |                |

| Id | Name | SBO | Value | Unit          | Constant |
|----|------|-----|-------|---------------|----------|
| s2 | s2   |     | 0.000 | dimensionless |          |

# 5 Rules

This is an overview of three rules.

#### **5.1** Rule x

Rule x is a rate rule for parameter x:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathbf{x} = \mathbf{c} \cdot \mathbf{y} - \mathbf{m}\mathbf{u}\mathbf{2} \cdot \mathbf{x} + \frac{\mathbf{p}\mathbf{1} \cdot \mathbf{x} \cdot \mathbf{z}}{\mathbf{g}\mathbf{1} + \mathbf{z}} + \mathbf{s}\mathbf{1} \tag{1}$$

**Derived unit** dimensionless

## **5.2 Rule** y

Rule y is a rate rule for parameter y:

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathbf{y} = \mathbf{r}2 \cdot \mathbf{y} \cdot (1 - \mathbf{b} \cdot \mathbf{y}) - \frac{\mathbf{a} \cdot \mathbf{x} \cdot \mathbf{y}}{\mathbf{g}2 + \mathbf{y}} \tag{2}$$

## **5.3 Rule** z

Rule z is a rate rule for parameter z:

$$\frac{\mathrm{d}}{\mathrm{d}t}z = \frac{\mathrm{p}2 \cdot \mathrm{x} \cdot \mathrm{y}}{\mathrm{g}3 + \mathrm{y}} - \mathrm{m}\mathrm{u}3 \cdot \mathrm{z} + \mathrm{s}2 \tag{3}$$

**Derived unit** dimensionless

SBML2ATEX 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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