## **SBML Model Report**

# Model name: "Grlich2003\_RanGTP\_gradient"



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### 1 General Overview

This is a document in SBML Level 2 Version 1 format. This model was created by the following three authors: Lukas Endler<sup>1</sup>, Vijayalakshmi Chelliah<sup>2</sup> and Dirk Gorlich<sup>3</sup> at September nineth 2008 at 2:04 p. m. and last time modified at April eighth 2016 at 3:46 p. m. Table 1 provides 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	13
events	0	constraints	0
reactions	9	function definitions	0
global parameters	0	unit definitions	5
rules	0	initial assignments	0

#### **Model Notes**

This model represents a concentration gradient of RanGTP across the nuclear envelope. This gradient is generated by distribution of regulators of RanGTPase. We have taken a log linear plot of graphs generated by GENESIS and compared with the experimental graphs.

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### 2 Unit Definitions

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

#### 2.1 Unit substance

**Definition** µmol

### 2.2 Unit microMpsec

Name microMpsec

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

### 2.3 Unit pmicroMpsec

Name pmicroMpsec

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

### 2.4 Unit psec

Name psec

**Definition**  $s^{-1}$ 

#### 2.5 Unit microM

Name microM

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

### 2.6 Unit volume

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

**Definition** 1

### 2.7 Unit area

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

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

### 2.8 Unit length

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

**Definition** m

## 2.9 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
nucleus cytoplasm			3 3	$1.2 \cdot 10^{-11} \\ 1.8 \cdot 10^{-11}$	1	<b>1</b>	

### 3.1 Compartment nucleus

This is a three dimensional compartment with a constant size of  $1.2 \cdot 10^{-11}$  litre.

### 3.2 Compartment cytoplasm

This is a three dimensional compartment with a constant size of  $1.8 \cdot 10^{-11}$  litre.

# 4 Species

This model contains 13 species. The boundary condition of two of these species is set to true so that these species' amount cannot be changed by any reaction. Section 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 Condi- tion
RanGDP_nuc		nucleus	$\mu$ mol·l <sup>-1</sup>		
$RCC1\_RanGDP$		nucleus	$\mu mol \cdot l^{-1}$		$\Box$
GDP		nucleus	$\mu mol \cdot l^{-1}$		
RCC1		nucleus	$\mu mol \cdot l^{-1}$		
$\tt RCC1\_RanGTP$		nucleus	$\mu mol \cdot l^{-1}$		
$RCC1\_Ran$		nucleus	$\mu mol \cdot l^{-1}$		
GTP		nucleus	$\mu mol \cdot l^{-1}$		
RanGTP_nuc		nucleus	$\mu mol \cdot l^{-1}$		
RanGAP		${ t cytoplasm}$	$\mu mol \cdot l^{-1}$		
RanBP1		${ t cytoplasm}$	$\mu mol \cdot l^{-1}$		
$RanGTP\_cy$		${ t cytoplasm}$	$\mu mol \cdot l^{-1}$		
$RanGTP\_RanBP1$		${ t cytoplasm}$	$\mu mol \cdot l^{-1}$		
$RanGDP_cy$		${ t cytoplasm}$	$\mu mol \cdot l^{-1}$		$\Box$

# **5 Reactions**

This model contains nine 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

N₀	Id	Name	Reaction Equation	SBO
1	RCC1_binding		$RanGDP\_nuc + RCC1 \Longrightarrow RCC1\_RanGDP$	
2	GDP-		$RCC1_RanGDP \Longrightarrow RCC1_Ran + GDP$	
	$\_$ dissociation			
3	$\mathtt{GTP\_binding}$		$RCC1\_Ran + GTP \Longrightarrow RCC1\_RanGTP$	
4	$RanGTP\_release$		$RCC1\_RanGTP \Longrightarrow RanGTP\_nuc + RCC1$	
5	Cytoplasmic-		RanGTP_nuc <del>←</del> RanGTP_cy	
	$_{ extsf{ extsf}}$ transfer			
6	Nucleoplasmic-		RanGDP_nuc <del>←</del> RanGDP_cy	
	$_{ t transfer}$			
7	${\tt RanGTP\_binding}$		$RanGTP\_cy + RanBP1 \Longrightarrow RanGTP\_RanBP1$	
8	RanBP1_RanGDP	RanBP1_RanGDP	$RanGTP\_RanBP1 \xrightarrow{RanGAP} RanGDP\_cy + RanBP1$	
9	${\tt RanGAP\_RanGDP}$	RanGAP_RanGDP	RanGTP_cy RanGDP_cy	

### **5.1 Reaction RCC1\_binding**

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

### **Reaction equation**

$$RanGDP\_nuc + RCC1 \Longrightarrow RCC1\_RanGDP \qquad (1)$$

#### **Reactants**

Table 5: Properties of each reactant.

Id	Name	SBO
RanGDP_nuc		
RCC1		

#### **Product**

Table 6: Properties of each product.

Id	Name	SBO
RCC1_RanGDP		

#### **Kinetic Law**

Derived unit  $s^{-1} \cdot \mu mol$ 

$$v_1 = \text{vol}(\text{nucleus}) \cdot (\text{r1} \cdot [\text{RanGDP\_nuc}] \cdot [\text{RCC1}] - \text{r8} \cdot [\text{RCC1\_RanGDP}])$$
 (2)

Table 7: Properties of each parameter.

Id	Name	SBO Val	lue Unit	Constant
r1	r1		$1 \cdot \mu \text{mol}^{-1} \cdot \text{s}^{-1}$	
r8	r8	55	$5.0   s^{-1}$	

### **5.2 Reaction** GDP\_dissociation

This is a reversible reaction of one reactant forming two products.

### **Reaction equation**

$$RCC1\_RanGDP \Longrightarrow RCC1\_Ran + GDP$$
 (3)

#### Reactant

Table 8: Properties of each reactant.

Id	Name	SBO
RCC1_RanGDP		

### **Products**

Table 9: Properties of each product.

Id	Name	SBO
RCC1_Ran		
GDP		

### **Kinetic Law**

Derived unit  $s^{-1} \cdot \mu mol$ 

$$v_2 = \text{vol}(\text{nucleus}) \cdot (\text{r2} \cdot [\text{RCC1\_RanGDP}] - \text{r7} \cdot [\text{RCC1\_Ran}] \cdot [\text{GDP}]) \tag{4}$$

Table 10: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
r2	r2	$21.0   s^{-1}$	
r7	r7	$11.0  1 \cdot \mu \text{mol}^{-1} \cdot \text{s}^{-1}$	$\square$

### **5.3 Reaction GTP\_binding**

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

### **Reaction equation**

$$RCC1\_Ran + GTP \Longrightarrow RCC1\_RanGTP$$
 (5)

### **Reactants**

Table 11: Properties of each reactant.

Id	Name	SBO
RCC1_Ran		
GTP		

### **Product**

Table 12: Properties of each product.

Id	Name	SBO
RCC1_RanGTP		

### **Kinetic Law**

Derived unit  $s^{-1} \cdot \mu mol$ 

$$v_3 = \text{vol}(\text{nucleus}) \cdot (\text{r3} \cdot [\text{RCC1\_Ran}] \cdot [\text{GTP}] - \text{r6} \cdot [\text{RCC1\_RanGTP}])$$
 (6)

Table 13: Properties of each parameter.

		•			
Id	Name	SBO	Value	Unit	Constant
r3 r6	r3 r6		0.6 19.0	$\frac{1 \cdot \mu mol^{-1} \cdot s^{-1}}{s^{-1}}$	

### **5.4 Reaction RanGTP\_release**

This is a reversible reaction of one reactant forming two products.

### **Reaction equation**

$$RCC1\_RanGTP \Longrightarrow RanGTP\_nuc + RCC1$$
 (7)

#### Reactant

Table 14: Properties of each reactant.

Id	Name	SBO
RCC1_RanGTP		

### **Products**

Table 15: Properties of each product.

Id	Name	SBO
RanGTP_nuc RCC1		

### **Kinetic Law**

Derived unit  $s^{-1} \cdot \mu mol$ 

$$\textit{v}_{4} = vol\left(nucleus\right) \cdot \left(r4 \cdot \left[RCC1\_RanGTP\right] - r5 \cdot \left[RanGTP\_nuc\right] \cdot \left[RCC1\right]\right) \tag{8}$$

Table 16: Properties of each parameter.

Id	Name	SBO Value	Unit	Constant
r4	r4	55.0	$s^{-1}$	$\overline{Z}$
r5	r5	100.0	$1 \cdot \mu \text{mol}^{-1} \cdot \text{s}^{-1}$	

# **5.5 Reaction** Cytoplasmic\_transfer

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

### **Reaction equation**

$$RanGTP\_nuc \rightleftharpoons RanGTP\_cy \tag{9}$$

### Reactant

Table 17: Properties of each reactant.

Id	Name	SBO
RanGTP_nuc		

#### **Product**

Table 18: Properties of each product.

Id	Name	SBO
RanGTP_cy		

### **Kinetic Law**

Derived unit  $s^{-1} \cdot \mu mol$ 

$$v_5 = \text{kpermRanGTP} \cdot \text{vol}(\text{nucleus}) \cdot ([\text{RanGTP\_nuc}] - [\text{RanGTP\_cy}])$$
 (10)

Table 19: Properties of each parameter.

Id	Name	SBO	Value	Unit	Constant
kpermRanGTP	kpermRanGTP		0.03	$s^{-1}$	

### **5.6 Reaction** Nucleoplasmic\_transfer

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

### **Reaction equation**

$$RanGDP\_nuc \rightleftharpoons RanGDP\_cy \tag{11}$$

### Reactant

Table 20: Properties of each reactant.

Id	Name	SBO
RanGDP_nuc		

### **Product**

Table 21: Properties of each product.

Id	Name	SBO
$RanGDP_cy$		

### **Kinetic Law**

Derived unit  $s^{-1} \cdot \mu mol$ 

$$v_6 = \text{kpermRanGDP} \cdot \text{vol}(\text{nucleus}) \cdot ([\text{RanGDP\_nuc}] - [\text{RanGDP\_cy}])$$
 (12)

Table 22: Properties of each parameter.

Id	Name	SBO	Value	Unit	Constant
kpermRanGDP	kpermRanGDP		0.12	$s^{-1}$	

### 5.7 Reaction RanGTP\_binding

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

### **Reaction equation**

$$RanGTP\_cy + RanBP1 \Longrightarrow RanGTP\_RanBP1$$
 (13)

#### **Reactants**

Table 23: Properties of each reactant.

Id	Name	SBO
RanGTP_cy		
RanBP1		

#### **Product**

Table 24: Properties of each product.

Id	Name	SBO
RanGTP_RanBP1		

#### **Kinetic Law**

**Derived unit**  $s^{-1} \cdot \mu mol$ 

$$v_7 = (\text{kon} \cdot [\text{RanGTP\_cy}] \cdot [\text{RanBP1}] - \text{koff} \cdot [\text{RanGTP\_RanBP1}]) \cdot \text{vol}(\text{cytoplasm})$$
 (14)

Table 25: Properties of each parameter.

Id	Name	SBO V	Value Unit	Constant
kon koff	kon koff		$0.300  1 \cdot \mu \text{mol} \\ \cdot 10^{-4}  \text{s}^{-1}$	-1·s <sup>-1</sup>

### 5.8 Reaction RanBP1\_RanGDP

This is a reversible reaction of one reactant forming two products influenced by one modifier.

Name RanBP1\_RanGDP

### **Reaction equation**

$$RanGTP\_RanBP1 \xrightarrow{RanGAP} RanGDP\_cy + RanBP1$$
 (15)

### Reactant

Table 26: Properties of each reactant.

Id	Name	SBO
RanGTP_RanBP1		

#### **Modifier**

Table 27: Properties of each modifier.

Id	Name	SBO
RanGAP		

#### **Products**

Table 28: Properties of each product.

Id	Name	SBO
RanGDP_cy		
RanBP1		

#### **Kinetic Law**

Derived unit  $s^{-1} \cdot 10^{-6} \text{ mol}$ 

$$v_8 = \frac{\text{vol}\left(\text{cytoplasm}\right) \cdot \text{kcat} \cdot [\text{RanGTP\_RanBP1}] \cdot [\text{RanGAP}]}{[\text{RanGTP\_RanBP1}] + \text{Km}} \tag{16}$$

Table 29: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
kcat	kcat	$10.8   s^{-1}$	$\square$
Km	Km	$0.1  \mu \text{mol} \cdot l^{-1}$	

### 5.9 Reaction RanGAP\_RanGDP

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

### Name RanGAP\_RanGDP

### **Reaction equation**

$$RanGTP\_cy \xrightarrow{RanGAP} RanGDP\_cy$$
 (17)

#### Reactant

Table 30: Properties of each reactant.

Id	Name	SBO
RanGTP_cy		

#### **Modifier**

Table 31: Properties of each modifier.

Id	Name	SBO
RanGAP		

### **Product**

Table 32: Properties of each product.

Id	Name	SBO
$RanGDP_cy$		

### **Kinetic Law**

Derived unit  $s^{-1} \cdot 10^{-6} \text{ mol}$ 

$$v_9 = \frac{\text{vol}\left(\text{cytoplasm}\right) \cdot \text{kcat\_GAP} \cdot [\text{RanGTP\_cy}] \cdot [\text{RanGAP}]}{\text{Km\_GAP} + [\text{RanGTP\_cy}]} \tag{18}$$

Table 33: Properties of each parameter.

Id	Name	SBO Value Unit	Constant
kcat_GAP Km_GAP	kcat Km	10.6 $s^{-1}$ 0.7 $\mu$ mol· $l^{-1}$	

# **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.

### 6.1 Species RanGDP\_nuc

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

This species takes part in two reactions (as a reactant in RCC1\_binding, Nucleoplasmic\_transfer).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{RanGDP\_nuc} = -v_1 - v_6 \tag{19}$$

### 6.2 Species RCC1\_RanGDP

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

This species takes part in two reactions (as a reactant in GDP\_dissociation and as a product in RCC1\_binding).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{RCC1}_{-}\mathrm{RanGDP} = v_1 - v_2 \tag{20}$$

### 6.3 Species GDP

Initial concentration  $1.6 \ \mu mol \cdot l^{-1}$ 

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

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{GDP} = 0\tag{21}$$

### **6.4 Species RCC1**

Initial concentration  $0.7~\mu mol \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in RCC1\_binding and as a product in RanGTP\_release).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{RCC1} = v_4 - v_1 \tag{22}$$

### 6.5 Species RCC1\_RanGTP

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

This species takes part in two reactions (as a reactant in RanGTP\_release and as a product in GTP\_binding).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{RCC1}_{-}\mathrm{RanGTP} = v_3 - v_4 \tag{23}$$

### 6.6 Species RCC1\_Ran

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

This species takes part in two reactions (as a reactant in GTP\_binding and as a product in GDP-dissociation).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{RCC1}_{-}\mathrm{Ran} = v_2 - v_3 \tag{24}$$

### 6.7 Species GTP

Initial concentration  $500 \ \mu mol \cdot l^{-1}$ 

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

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{GTP} = 0\tag{25}$$

### 6.8 Species RanGTP\_nuc

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

This species takes part in two reactions (as a reactant in Cytoplasmic\_transfer and as a product in RanGTP\_release).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{RanGTP\_nuc} = v_4 - v_5 \tag{26}$$

### 6.9 Species RanGAP

Initial concentration  $0.7 \, \mu mol \cdot l^{-1}$ 

This species takes part in two reactions (as a modifier in RanBP1\_RanGDP, RanGAP\_RanGDP).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{RanGAP} = 0\tag{27}$$

### 6.10 Species RanBP1

Initial concentration  $2 \mu mol \cdot l^{-1}$ 

This species takes part in two reactions (as a reactant in RanGTP\_binding and as a product in RanBP1\_RanGDP).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{RanBP1} = v_8 - v_7 \tag{28}$$

#### **6.11 Species RanGTP\_cy**

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

This species takes part in three reactions (as a reactant in RanGTP\_binding, RanGAP\_RanGDP and as a product in Cytoplasmic\_transfer).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{RanGTP}_{-}\mathrm{cy} = v_5 - v_7 - v_9 \tag{29}$$

### 6.12 Species RanGTP\_RanBP1

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

This species takes part in two reactions (as a reactant in RanBP1\_RanGDP and as a product in RanGTP\_binding).

$$\frac{d}{dt}RanGTP\_RanBP1 = v_7 - v_8 \tag{30}$$

### **6.13 Species** RanGDP\_cy

Initial concentration 5 µmol·l<sup>-1</sup>

This species takes part in three reactions (as a product in Nucleoplasmic\_transfer, RanBP1-\_RanGDP, RanGAP\_RanGDP).

$$\frac{\mathrm{d}}{\mathrm{d}t}\mathrm{RanGDP}_{-}\mathrm{cy} = v_6 + v_8 + v_9 \tag{31}$$

BML2ATEX 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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