

# Long-time behaviour of the stochastic models of stem cells differentiation.

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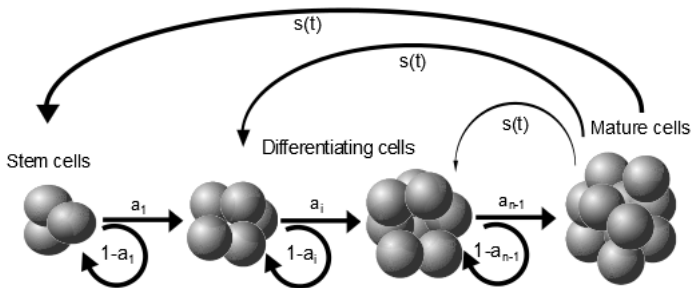


Figure: The process of uni-potential stem cells differentiation

$$\begin{aligned}\frac{dc_1}{dt} &= (2a_1s_{k_1}(t) - 1)p_1c_1(t) - \mu_1c_1, \\ \frac{dc_2}{dt} &= (2a_2s_{k_1}(t) - 1)p_2c_2(t) + 2(1 - a_1s_{k_1}(t))p_1c_1(t) - \mu_2c_2(t), \\ &\vdots \\ \frac{dc_n}{dt} &= 2(1 - a_{n-1}s_{k_1}(t))p_{n-1}c_{n-1}(t) - \mu_nc_n(t),\end{aligned}\tag{1}$$

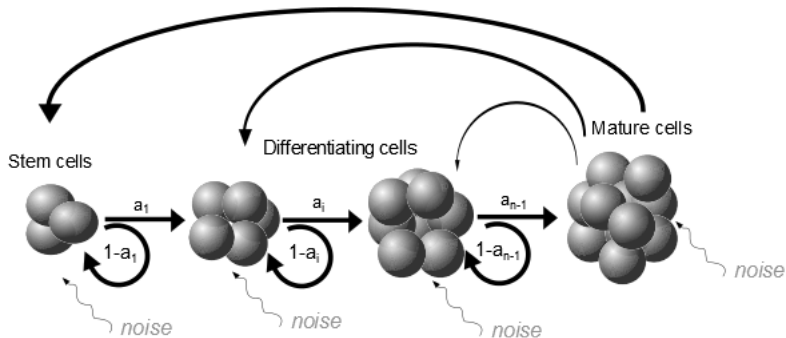


Figure: The process of uni-potential stem cells differentiation with noise

$$\begin{aligned}
 dc_1 &= \left( \left( \frac{2a_1}{1+kc_n} - 1 \right) p_1 c_1 - \mu_1 c_1 \right) dt \\
 dc_2 &= \left( \left( \frac{2a_2}{1+kc_n} - 1 \right) p_2 c_2 \right. \\
 &\quad \left. + 2 \left( 1 - \frac{a_1}{1+kc_n} \right) p c_1 - \mu_2 c_2 \right) dt \\
 &\quad \vdots \\
 dc_n &= \left( 2 \left( 1 - \frac{a_{n-1}}{1+kc_n} \right) p_{n-1} c_{n-1} - \mu_n c_n \right) dt
 \end{aligned} \tag{2}$$

$$dc_1 = \left( \left( \frac{2a_1}{1+kc_n} - 1 \right) p_1 c_1 - \mu_1 c_1 \right) dt + \alpha_1 c_1 dW_{1,t}$$

$$dc_2 = \left( \left( \frac{2a_2}{1+kc_n} - 1 \right) p_2 c_2 \right. \\ \left. + 2 \left( 1 - \frac{a_1}{1+kc_n} \right) p c_1 - \mu_2 c_2 \right) dt + \alpha_2 c_2 dW_{2,t}$$

$$\vdots$$

$$dc_n = \left( 2 \left( 1 - \frac{a_{n-1}}{1+kc_n} \right) p_{n-1} c_{n-1} - \mu_n c_n \right) dt + \alpha_n c_n dW_{n,t} \quad (2)$$

$$\begin{aligned}dc_1 &= \left( \frac{2a}{1+kc_n} - 1 \right) pc_1 dt + \alpha_1 c_1 dW_{1,t} \\dc_2 &= \left( 2 \left( 1 - \frac{a}{1+kc_2} \right) pc_1 - \mu c_2 \right) dt + \alpha_2 c_2 dW_{2,t} \quad (3)\end{aligned}$$

Where  $\alpha_1, \alpha_2 > 0$ .

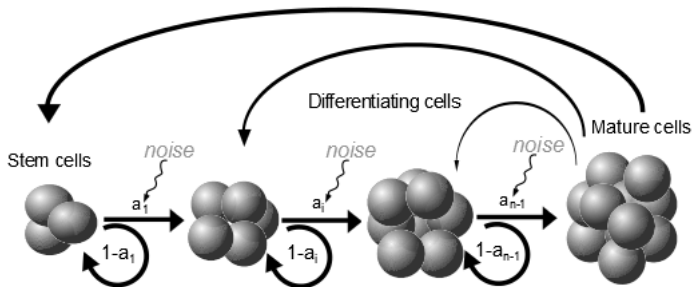
## Theorem

*Let the following condition*

$$(2a - 1)p - \frac{\alpha_1^2}{2} > 0 \quad (4)$$

*holds, then the semi-group  $\{P(t)\}_{t \geq 0}$  related to the model (??) is asymptotically stable.*





**Figure:** The process of uni-potential stem cells differentiation with the perturbation of the fraction of self-renewal

$$\begin{aligned}x_1'(t) &= \left( \frac{2a}{1+kx_2(t)} - 1 \right) px_1(t) \\x_2'(t) &= 2 \left( 1 - \frac{a}{1+kx_2(t)} \right) px_1(t) - dx_2(t)\end{aligned}\quad (5)$$

$$\begin{aligned}x_1'(t) &= \left( \frac{2a_t}{1+kx_2(t)} - 1 \right) px_1(t) \\x_2'(t) &= 2 \left( 1 - \frac{a_t}{1+kx_2(t)} \right) px_1(t) - dx_2(t)\end{aligned}\quad (5)$$

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$$a_t = \{a_0, a_1\}, \quad 0 < a_0 < \frac{1}{2} < a_1 < 1 \quad (6)$$

$$\begin{aligned}x_1'(t) &= \left( \frac{2a_t}{1+kx_2(t)} - 1 \right) px_1(t) \\x_2'(t) &= 2 \left( 1 - \frac{a_t}{1+kx_2(t)} \right) px_1(t) - dx_2(t)\end{aligned}\quad (5)$$

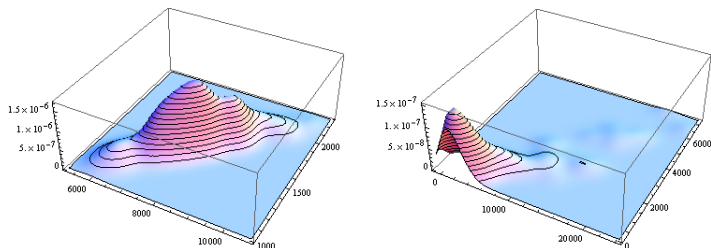
$$a_t = \{a_0, a_1\}, \quad 0 < a_0 < \frac{1}{2} < a_1 < 1 \quad (6)$$

## Theorem

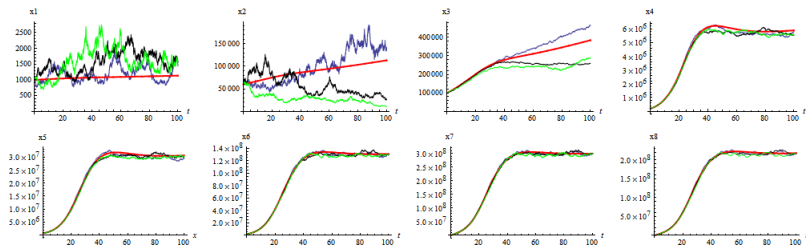
Let  $\{P(t)\}_{t \geq 0}$  be the semi-group related to the model (5) and let the following condition

$$\frac{2a_0 - 1}{1 - 2a_1} > \frac{q_0}{q_1} \quad (7)$$

holds. Then the semi-group  $\{P(t)\}_{t \geq 0}$  is asymptotically stable.

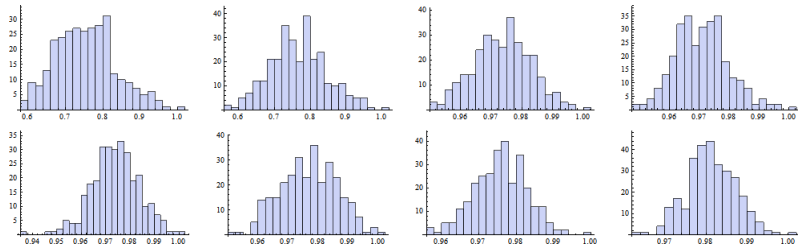


**Figure:** Densities of the process (3) for  $T_{end} = 500$ ,  $p = 0.6$ ,  $\mu = 2.77$ ,  $k = 1, 28 \cdot 10^{-6}$  and two sets of  $\alpha$  and  $a$ : ,  $a = 0.6$ ,  $\alpha = 0.1$ ,  $a = 0.3$ ,  $\alpha = 1$

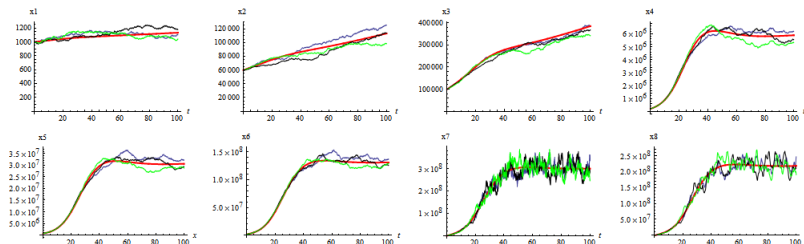


**Figure:** A solution of the model (2) and 3 trajectories (blue, green, black) of the stochastic processes in time  $T_{end} = 100$  of the model (??) for  $n = 8$  ( $\alpha_1 = 1, \alpha_2 = 1, \alpha_3 = 1, \alpha_4 = 0.017, \alpha_5 = 0.019, \alpha_6 = 0.1, \alpha_7 = 0.1, \alpha_8 = 0.1$ ) and  $a_1 = 0.735, a_2 = 0.7298, a_3 = 0.7245, a_4 = 0.7140, a_5 = 0.5775, a_6 = 0.4725, a_7 = 0.3675, \rho_1 = 0.006, \rho_2 = 0.03, \rho_3 = 0.18, \rho_4 = 0.6, \rho_5 = 0.65, \rho_6 = 1, \rho_7 = 1.5, \mu_1 = \mu_2 = \mu_3 = \mu_4 = \mu_5 = \mu_6 = \mu_7 = 0, \mu_8 = 2.77, k = 1,28 \cdot 10^{-9}$ .

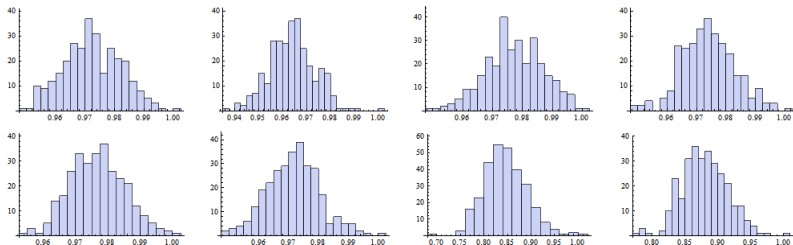




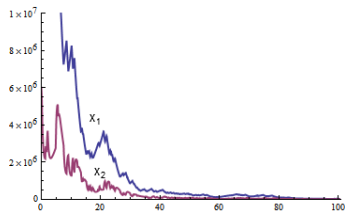
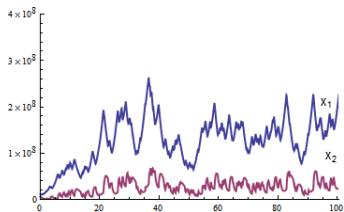
**Figure:** Histograms for 300 trajectories for a time  $T_{end} = 200$  of the model (??) for  $n = 8$  ( $\alpha_1 = 1, \alpha_2 = 1, \alpha_3 = 1, \alpha_4 = 0.017, \alpha_5 = 0.019, \alpha_6 = 0.1, \alpha_7 = 0.1, \alpha_8 = 0.1$ ) and  $a_1 = 0.735, a_2 = 0.7298, a_3 = 0.7245, a_4 = 0.7140, a_5 = 0.5775, a_6 = 0.4725, a_7 = 0.3675, p_1 = 0.006, p_2 = 0.03, p_3 = 0.18, p_4 = 0.6, p_5 = 0.65, p_6 = 1, p_7 = 1.5, \mu_1 = \mu_2 = \mu_3 = \mu_4 = \mu_5 = \mu_6 = \mu_7 = 0, \mu_8 = 2.77, k = 1,28 \cdot 10^{-9}$ .



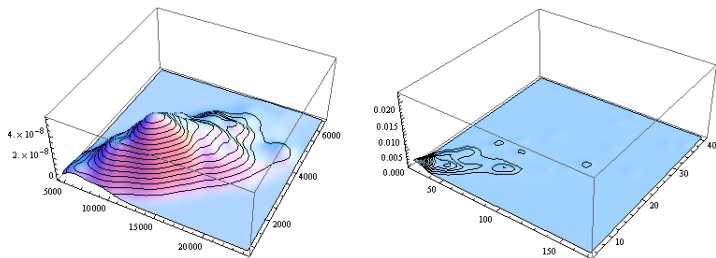
**Figure:** A solution of the model (2) and 3 trajectories (blue, green, black) of stochastic processes in time  $T_{end} = 100$  of the model (??) for  $n = 8$  with 2 ( $\alpha_1 = 0.1$ ,  $\alpha_2 = 0.1$ ,  $\alpha_3 = 0.1$ ,  $\alpha_4 = 0.017$ ,  $\alpha_5 = 0.015$ ,  $\alpha_6 = 0.12$ ,  $\alpha_7 = 1$ ,  $\alpha_8 = 1$ ) and  $a_1 = 0.735$ ,  $a_2 = 0.7298$ ,  $a_3 = 0.7245$ ,  $a_4 = 0.7140$ ,  $a_5 = 0.5775$ ,  $a_6 = 0.4725$ ,  $a_7 = 0.3675$ ,  $p_1 = 0.006$ ,  $p_2 = 0.03$ ,  $p_3 = 0.18$ ,  $p_4 = 0.6$ ,  $p_5 = 0.65$ ,  $p_6 = 1$ ,  $p_7 = 1.5$ ,  $\mu_1 = \mu_2 = \mu_3 = \mu_4 = \mu_5 = \mu_6 = \mu_7 = 0$ ,  $\mu_8 = 2.77$ ,  $k = 1,28 \cdot 10^{-9}$ .



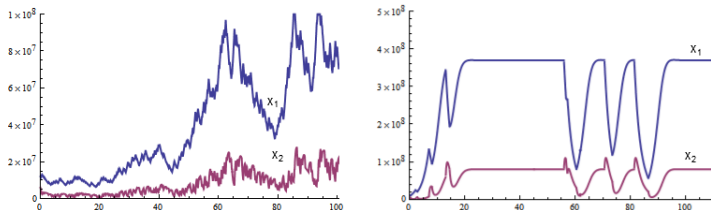
**Figure:** Histograms for 300 trajectories for a time  $T_{end} = 200$  of the model (??) for  $n = 8$  with 2 ( $\alpha_1 = 0.1$ ,  $\alpha_2 = 0.1$ ,  $\alpha_3 = 0.1$ ,  $\alpha_4 = 0.017$ ,  $\alpha_5 = 0.015$ ,  $\alpha_6 = 0.12$ ,  $\alpha_7 = 1$ ,  $\alpha_8 = 1$ ) and  $a_1 = 0.735$ ,  $a_2 = 0.7298$ ,  $a_3 = 0.7245$ ,  $a_4 = 0.7140$ ,  $a_5 = 0.5775$ ,  $a_6 = 0.4725$ ,  $a_7 = 0.3675$ ,  $p_1 = 0.006$ ,  $p_2 = 0.03$ ,  $p_3 = 0.18$ ,  $p_4 = 0.6$ ,  $p_5 = 0.65$ ,  $p_6 = 1$ ,  $p_7 = 1.5$ ,  $\mu_1 = \mu_2 = \mu_3 = \mu_4 = \mu_5 = \mu_6 = \mu_7 = 0$ ,  $\mu_8 = 2.77$ ,  $k = 1,28 \cdot 10^{-9}$ .



**Figure:** Trajectories of the processes of the model (5) for  $T_{end} = 100$ ,  $p = 0.6$ ,  $\mu = 2.77$ ,  $k = 1,28 \cdot 10^{-6}$ ,  $q_2 = 3$ ,  $q_1 = 1$  and  $a_2 = 0.3$ ,  $a_1 = 0.9$  (left one),  $a_2 = 0.2$ ,  $a_1 = 0.6$  (right one)



**Figure:** Density of the process (5) for  $T_{end} = 150$  and two different sets of parameters:  $a_1 = 0.8$ ,  $a_2 = 0.3$  and  $a_1 = 0.6$ ,  $a_2 = 0.2$ ,  $p = 0.6$ ,  $\mu = 2.77$ ,  $k = 1,28 \cdot 10^{-4}$ .



**Figure:** Trajectories of the processes of the model (5) for  $T_{end} = 100$ ,  $\rho = 0.6$ ,  $\mu = 2.77$ ,  $k = 1,28 \cdot 10^{-6}$ ,  $a_2 = 0.3$ ,  $a_1 = 0.9$  and  $q_0 = 4$ ,  $q_1 = 6$  (left one),  $q_0 = 0.3$ ,  $q_1 = 0.1$  (right one)



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# Thank you.