## A M system modelling membrane formation, cytoskeleton growth and reproduction

The M system described in this section is inspired by the process of mitosis and cell replication, controlled by the growth of cytoskeleton, as illustrated schematically at Fig. 2. Observe that, even if this process relies on several relatively complex regulation mechanisms, the M system model can simulate it, at least at the morphological and dynamical level, with as few as 16 rules. All processes in the model are controlled by fully local interactions between its elements.



Fig. 2. Major events in mitosis. Licensed under Wikimedia Commons, authored by Mysid.

Let us start the formal description of the model by its polytopic tile system  $T_0 = (Q, G, \gamma, d_g, S)$  in  $\mathbb{R}^3$ , which determines its spatial structure. Let

- $Q = \{q_0, q_1, q_2, q_3, q_4, s_0, s_1, s_2\}$ , where  $q_0, q_1, q_2$  are larger pentagonal tiles forming cellular membrane, and  $q_3, q_4$  are small pentagonal tiles forming nuclear membrane. All tiles contain facet connectors at all edges for mutual interconnection. There are three rods:  $s_0$  and  $s_1$  with two connectors at their endpoints, and  $s_2$  is a rod with one connector at one endpoint and two fork-oriented connectors at the other endpoint. Rod  $s_0$  is used only in the first step of the system. Rods  $s_1$  and  $s_2$  form straight and fork segments of cytoskeleton, respectively, and we call them *microtubules*. Furthermore,
  - tile  $q_0$  contains at the centre a point connector to which rod  $s_0$  can connect by its endpoint connector;
  - tile tile  $q_3$  contains at the centre a point connector which can connect to the other end of rod  $s_0$ ;
  - tiles  $q_0$  and  $q_1$  contain at the centre a point connector to which rod  $s_1$  can connect.

Let

 $\Delta_p$  be a regular pentagon with distance 10 from center to vertices, with vertices denoted by  $\mathbf{x}_1, \ldots, \mathbf{x}_5$ ;

 $\Delta'_p$  be a regular pentagon with distance 3 from center to vertices, with vertices denoted by  $\mathbf{x}'_1, \ldots, \mathbf{x}'_5$ ;

 $\begin{array}{l} \varDelta_0 \ = \langle 0, 9.184 \rangle \mbox{ be a segment with length } 9.184; \\ \varDelta_s \ = \langle 0, 1.4 \rangle \mbox{ be a segment with length } 1.4. \end{array}$ 

The tiles in Q are defined as follows:

 $q_0 = (\Delta_p,$ (pentagon)  $\{(\langle \mathbf{x}_i, \mathbf{x}_i \mod 5+1 \rangle, g_e, \varphi_p) \mid 1 \le i \le 5\}$ (facet connectors)  $\cup \{ (\mathbf{0}, g_0, \pi/2), (\mathbf{0}, g_3, \pi/2) \},\$ (point connectors) (surface glue)  $g_x),$  $q_1 = (\Delta_p,$  $\{(\langle \mathbf{x}_i, \mathbf{x}_i \mod 5+1 \rangle, g_a, \varphi_p) \mid 1 \le i \le 5\}$  $\cup$  {(**0**,  $g_3, \pi/2$ )},  $g_x$ ),  $q_2 = (\Delta_p,$  $\{(\langle \mathbf{x}_1, \mathbf{x}_2 \rangle, g_f, \varphi_p), (\langle \mathbf{x}_2, \mathbf{x}_3 \rangle, g_b, \varphi_p), (\langle \mathbf{x}_3, \mathbf{x}_4 \rangle, g_c, \varphi_p), (\langle \mathbf{x}_4, \mathbf{x}_5 \rangle, g_d, \varphi$  $(\langle \mathbf{x}_5, \mathbf{x}_1 \rangle, g_b, \varphi_p) \}, g_x),$  $q_3 = (\Delta'_p, \{(\langle \mathbf{x}'_i, \mathbf{x}'_i \mod 5+1 \rangle, g_a, \varphi_p) \mid 1 \le i \le 5\}$  $\cup \{(\mathbf{0}, g_2, -\pi/2),$  $g_t),$  $q_4 = (\Delta'_p,$  $\{(\langle \mathbf{x}_1', \mathbf{x}_2' \rangle, g_f, \varphi_p), (\langle \mathbf{x}_2', \mathbf{x}_3' \rangle, g_b, \varphi_p), (\langle \mathbf{x}_3', \mathbf{x}_4' \rangle, g_c, \varphi_p), (\langle \mathbf{x}_4', \mathbf{x}_5' \rangle, g_d, \varphi_p), (\langle$  $(\langle \mathbf{x}_5', \mathbf{x}_1' \rangle, g_b, \varphi_p) \}, g_x),$  $s_0 = (\Delta_0, \{(\{0\}, g_1, \pi/2), (\{9.184\}, g_1, \pi/2)\}, g_x);$  $s_1 = (\Delta_s, \{(\{0\}, g_3, 0), (\{1.4\}, g_4, 0)\}, g_x);$  $s_2 = (\Delta_s, \{(\{0\}, g_5, 0), (\{1.4\}, g_6, \pi/30), (\{2\}, g_4, -\pi/30)\}, g_x);$ where  $\varphi_p = 2.034443935795703$  rad is the inner angle between two faces of a dodecahedron.

 $\begin{aligned} G &= \{g_0, g_1, g_2, g_3, g_4, g_5, g_6, g_a, g_b, g_c, g_d, g_e, g_f, g_t, g_x\} \text{ is the set of glues;} \\ \gamma &= \{(g_0, g_1), (g_1, g_2), (g_3, g_3), (g_4, g_5), (g_6, g_3), (g_4, g_t), (g_6, g_t), (g_a, g_f), (g_f, g_a), \\ &\quad (g_b, g_b), (g_c, g_c), (g_d, g_d), (g_e, g_f)\} \text{ is the glue relation;} \\ d_g &= 0.1 \text{ is the glue distance;} \end{aligned}$ 

 $S = \{q_0\}$  is the seed tile.

Then let us define the M system  $\mathcal{M}_0 = (F, P, T_0, \mu, R, r, \sigma)$  such that:

$$\begin{split} F &= (O,m,\epsilon), \text{ where:} \\ & O = \{a,b,c,x\} \text{ are floating objects;} \\ & m(a) = 5, \ m(b) = 5, \ m(c) = 4, \ m(x) = 7 \text{ is the mobility of floating objects;} \\ & \epsilon(a) = 0.1 \text{ and } \epsilon(o) = 0 \text{ for all other } o \in O \text{ is the concentration of floating objects in the environment;} \end{split}$$

$$P = \{p_0, p_1, p_2, p_3, p_4\};$$

 $T_0$  is the polytopic tile system described above;

$$\begin{split} \mu(q_2) &= \{(p_0,(2,0)), (p_0,(2,2)), (p_2,(0,0)), (p_2,(0,2)), \} \cup \{(p_1,(i,j)) \mid i,j \in \{-5,5\}\}, \quad \text{(proteins placed on tiles)} \\ \mu(q_4) &= \{(p_3,(i,j)) \mid i,j \in \{-1,1\}\} \cup \{(p_4,(i,j)) \mid i,j \in \{0,1\}\}, \\ \mu(t) &= \emptyset \text{ for all other } t \in Q; \\ R \text{ contains the following rules:} \end{split}$$

Metabolic rules:  $a[p_0 \rightarrow [p_0 a;$   $[p_1 c \rightarrow [p_1 a;$   $[p_1 c \rightarrow [p_1 a;$   $[p_2 a \rightarrow [p_2 b;$   $[p_3 c x \rightarrow [p_3 aa;$   $cx[p_3 \rightarrow aa[p_3;$  $[p_4 a \rightarrow [p_4 cc;$ 

Creation rules:  $a^8 \rightarrow q_1$ ; (cellular membrane tiles creation from eight objects a)  $a^8 \rightarrow q_2$ ;  $aaa \rightarrow q_3$ ; (nuclear membrane tiles creation from three objects a)  $aaa \rightarrow q_4$ ;  $aaa \rightarrow s_0$ ; (auxiliary rod creation)  $b \rightarrow s_1$ ; (microtubules creation)  $b \rightarrow s_2$ ;

Division rules:

 $\begin{array}{l} g_c \xrightarrow{x} g_c \to g_c, g_c \quad \ (\text{division of both cellular and nuclear membrane}) \\ g_d \xrightarrow{x} g_d \to g_d, g_d \end{array}$ 

r = 14 is the reaction distance;

 $\sigma(g_4, g_t) = xxx$  and  $\sigma(g, h) = \emptyset$  for all other  $(g, h) \in \gamma$ .

The M system  $\mathcal{M}_0$  passes a (possibly infinite) sequence of configurations described below. Since the system is nondeterministic, the provided description captures its most probable development, with possible statistical deviations:

- 1. In the initial configuration there is a single seed tile  $q_0$  for cellular membrane. There are objects a in the environment in concentration 0.1 per cubic unit.
- 2. In the first step, the auxiliary rod  $s_0$  is created and attached to the centre of the seed tile  $q_0$  so that it is perpendicular to it. Simultaneously, five cellular membrane tiles  $q_2$  are created by the rule  $a^8 \rightarrow q_2$ , and attached to five edge connectors of the tile  $q_0$ . Their other connectors at mutually matching positions with glues  $g_b$  attach together, too.
- 3. In the second step, tile  $q_3$  is created and attached perpendicularly to rod  $s_0$ , hence it is parallel with  $q_0$ . Simultaneously, five more tiles  $q_2$  are connected to the existing tiles  $q_2$ , forming subsequently the closed structure playing the role of cellular membrane.

- 4. In the third step, pentagonal tile  $q_1$  concludes the cellular membrane building phase, completing the dodecahedron-shaped cell ("soccer ball") and enclosing inside rather large number of objects a (over a hundred of thousands). Simultaneously, an analogous process of membrane creation on a smaller scale starts from tile  $q_3$  to which five tiles  $q_4$  are attached, and the nuclear membrane formation is completed in next two steps.
- 5. During the previous two steps, the reaction proteins  $p_2$  on tile  $q_2$  already started to catalyze the reaction of objects a to b applying the rule  $p_2a \rightarrow p_2b$ . As each tile  $q_2$  contains two proteins, and the cellular membrane contains 10 tiles  $q_2$ , 20 objects b is produced at each step within the cell. An analogous process runs in the nuclear membrane with proteins on tiles  $q_4$  and objects c.
- 6. The objects b allow for creation of the microtubules  $s_1$  and  $s_2$ , which can connect to point connectors in centres of "polar" tiles  $q_0$  and  $q_1$  of the cellular membrane. A non-deterministic growth of cytoskeleton towards the interior of the cell starts, consuming objects b as it continues.
- 7. When the microtubules  $s_1$  and  $s_2$  of the growing cytoskeleton reach the nuclear membrane, they can attach to its polar tiles  $q_3$ , each such contact releasing three objects x (as defined in the mapping  $\sigma$ ).
- 8. Each object x can break one bond of a pair of tiles  $q_2$  (or  $q_4$ ) in the equatorial part of both membranes.
- 9. When all these bonds (10 in each membrane) are broken, the cell splits into two parts, each containing a half of the outer membrane, a part of the cytoskeleton growing from it, and the corresponding half of the nucleus.
- 10. The same growth rules as in the second and the third step now allow to complete both halves into two complete cells, pushing them apart as they grow. Before concluding this process, however, two important events take place: (i) both open halves of the cellular membrane are replenished with objects *a* from the environment, and (ii) open nucleus releases a large amount of accumulated objects *c*, while replenishing *a*'s. Objects *c* diffuse into both halves of the cell, deactivating objects *x* via the rules  $[p_1cx \rightarrow [p_1aa, [p_3cx \rightarrow [p_3aa \text{ and } cx[p_3 \rightarrow aa[p_3.]])$
- 11. After two steps during which the two new cells are completed, cytoskeleton growth can continue and the process of the "cell" division is repeated.

In such a way, the cells in the M system continue their division and the whole system is in a dynamical homeostasis: if environmental resources (represented by objects a) are unlimited, then so is the growth of the system. In the limited environment the growth would stop after exhaustion of the environmental resources.

Notice that, as the model is nondeterministic, the mitosis can be sometimes blocked, for instance because the objects x regulating the division process may start another nucleus division before the whole cell is ready to divide. This synchronization disorder in division control makes the cell dysfunctional and incapable of further mitosis.