CEMRACS 2018 project: Mathematical modelling of cell aggregation and segregation.

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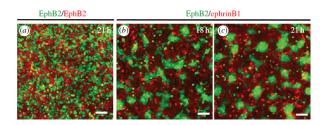
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Biological context

Cell segregation and border sharpening in two-species systems



Working hypothesis: inter(heterotypic) and intra(homotypic) species repulsion control cell segregation and border sharpening. They have more influence than inter- or intra- species adhesion.

Goal: to understand the mechanisms of morphogenesis.



blabla



blabla



Microscopic framework

Individual Based Model for particles interacting through repulsion interactions:

$$\begin{cases} dX_i^A = -\mu \nabla_{X_i^A} W^A(X^A, X^B) dt + \sqrt{2D_A} dB_i, & \forall i \in \{1, \dots, N_A\} \\ dX_i^B = -\mu \nabla_{X_\ell^A} W^B(X^A, X^B) dt + \sqrt{2D_B} dB_\ell, & \forall \ell \in \{1, \dots, N_B\} \end{cases}$$
(1)

where $\mu > 0$ is the mobility coefficient and B_i is a 2-dimensional Brownian motion $B_i = (B_i^1, B_i^2)$ of intensity $D_A, D_B > 0$ respectively for species A and B.





Macroscopic framework

$$\begin{cases}
\partial_{t}f^{A} = \nabla \cdot \underbrace{\left(f^{A}\nabla_{x}(\Phi^{AA} * f^{A}) + f^{A}\nabla_{x}(\Phi^{AB} * f^{B})\right)}_{interaction \ potential} + \underbrace{D_{A}\Delta_{x}f^{A}}_{diffusion} + \underbrace{\nu^{A}f^{A}\left(1 - \frac{f^{A} + f^{B}}{f^{*}}\right)}_{logistic \ term} \\
\partial_{t}f^{B} = \nabla \cdot \left(f^{B}\nabla_{x}(\Phi^{BB} * f^{B}) + f^{B}\nabla_{x}(\Phi^{BA} * f^{A})\right) + D_{A}\Delta_{x}f^{A} + \nu^{B}f^{B}\left(1 - \frac{f^{A} + f^{B}}{f^{*}}\right)
\end{cases}$$
(2)

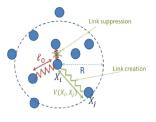
with $f^{S}(x,t)$ particle distributions of type S that give the probability to find a particle of type S at a point x and time t, $S \in \{A, B\}$.

 f^* : carrying capacity, it represents the maximum population size that can be present in the environment.



Macroscopic framework

We suppose that the homo-heterotypic species links act as a springs of rest length R between the particles that it is also detection radius for the interaction.



Case of Hookean springs

$$\Phi^{ST}(x) = \frac{\nu_c^{ST}}{\nu_d^{ST}} \frac{\kappa^{ST}}{2} \begin{cases} (|x| - R)^2, & \text{for } |x| \le R \\ 0, & \text{for } |x| > R \end{cases}$$

with ν_c^{ST}, ν_d^{ST} Poisson processes frequencies and κ^{ST} interaction intensity.

pр



blabla



Analysis of the macroscopic model

$$\begin{cases} \partial_t f^A = \nabla \cdot (f^A \nabla_x (\Phi^{AA} * f^A) + f^A \nabla_x (\Phi^{AB} * f^B)) + D_A \Delta_x f^A + \nu^A f^A \left(1 - \frac{f^A + f^B}{f^*} \right) \\ \partial_t f^B = \nabla \cdot (f^B \nabla_x (\Phi^{BB} * f^B) + f^B \nabla_x (\Phi^{BA} * f^A)) + D_A \Delta_x f^A + \nu^B f^B \left(1 - \frac{f^A + f^B}{f^*} \right) \end{cases}$$
(3)

Linearization around constant steady states \bar{f}^A, \bar{f}^B and Fourier transform:

$$\partial_{t} \begin{pmatrix} \hat{f}^{A} \\ \hat{f}^{B} \end{pmatrix} = \underbrace{\begin{pmatrix} -|y|^{2} (2\pi \bar{f}^{A} \hat{\Phi}^{AA}(y) + D_{A}) - \nu_{b}^{A} \frac{\bar{f}^{A}}{f^{*}} & -|y|^{2} 2\pi \bar{f}^{A} \hat{\Phi}^{AB}(y) - \nu_{b}^{A} \frac{\bar{f}^{A}}{f^{*}} \\ -|y|^{2} \bar{f}^{B} \hat{\Phi}^{BA}(y) - \nu_{b}^{B} \frac{\bar{f}^{B}}{f^{*}} & -|y|^{2} (2\pi \bar{f}^{B} \hat{\Phi}^{BB}(y) + D_{B}) - \nu_{b}^{B} \frac{\bar{f}^{B}}{f^{*}} \end{pmatrix}}_{M(y)} \begin{pmatrix} \hat{f}^{A} \\ \hat{f}^{B} \end{pmatrix}$$
(4)

The constant steady states will be unstable if:





Logistic vs. no logistic

Logistic growth model

No logistic model



Numerical simulations

Micro Macro

