

Kinetic models for chemotaxis

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Main Goal

- To study two different levels of description for chemotaxis and to study how these two levels are related.

General Picture

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Question:

Which is the set of equations that Φ obey?

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If

$$\Phi(t) = \lim_{\varepsilon \rightarrow 0} \Phi_\varepsilon(t), \quad t < T$$

(in some sense) then \mathcal{M} is the limit model of \mathcal{M}_ε .

Chemotaxis

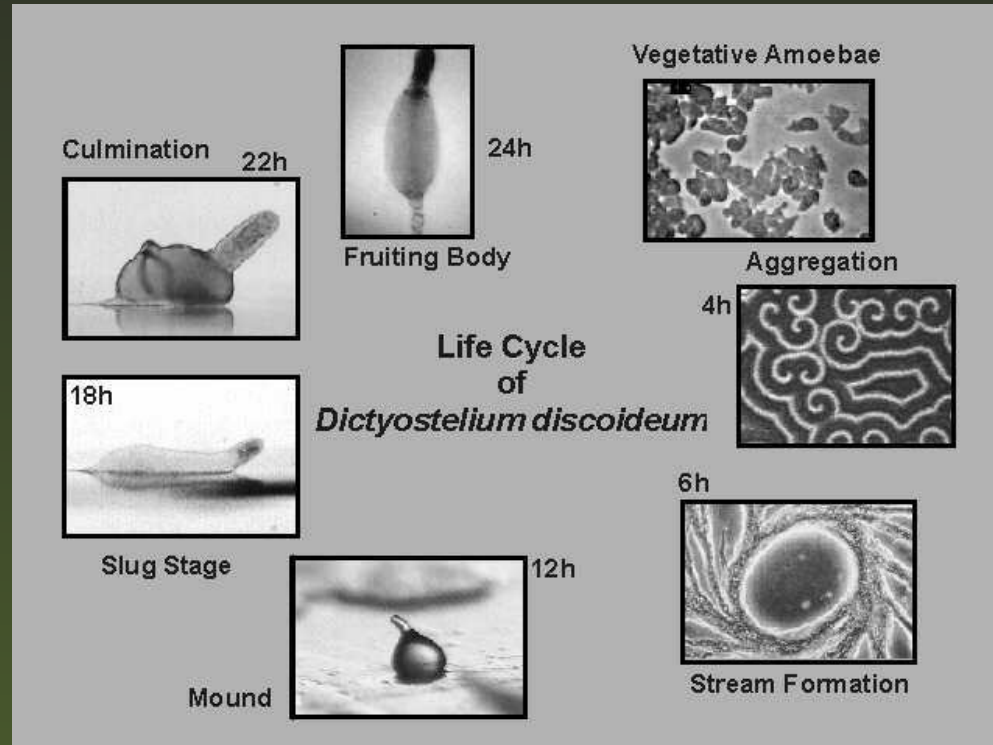


Figure 1: Life cycle of *Dictyostelium discoideum*. Picture made by Florian Seigert and Kees Wiejer (Zoologisches Institut München Ludwig-Maximilians-Universität München).

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- The cell goes in straight line for a certain characteristic time and then changes its direction from v' to v (in a space-time point (x, t) in the presence of the substance S and cell density ρ) according to a certain turning kernel $T[S, \rho](x, v, v', t)$.

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- The set of all possible velocities is given by a compact, spherically symmetric set V .

Velocity jump model

$$\partial_t f(x, v, t) + v \cdot \nabla f(x, v, t) = \int_V (T[S, \rho](x, v, v', t) f(x, v', t) - T[S, \rho](x, v', v, t) f(x, v, t)) dv' .$$

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$f(x, v, t)$ = cell density in space-time point (x, t) with velocity v (*phase-space density*).

Velocity jump model

Notation

$$f = f(x, v, t) ,$$

$$f' = f(x, v', t) ,$$

$$T[S, \rho] = T[S, \rho](x, v, v', t) ,$$

$$T^*[S, \rho] = T[S, \rho](x, v', v, t).$$

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Equation

$$\partial_t f + v \cdot \nabla f = \int_V (T[S, \rho] f' - T^*[S, \rho] f) dv' .$$

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- We should consider also an equation for S :

$$\partial_t S = D_0 \Delta S + \varphi(S, \rho) .$$

Keller-Segel model

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Keller-Segel equations:

$$\partial_t \rho = \nabla \cdot (D \nabla \rho - \chi(S) \beta(\rho) \rho \nabla S) ,$$

$$\partial_t S = D_0 \Delta S + \varphi(S, \rho) .$$

ρ = cell density ,

S = density of chemo-attractant ,

χ = chemotactic sensitivity ,

D, D_0 = diffusion coefficients ,

φ = interaction between ρ and S .

Keller-Segel model

- Typical example of interaction

$$\varphi(S, \rho) = \alpha\rho - \beta S ,$$

with $\alpha > 0, \beta \geq 0$.

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$$\varphi(S, \rho) = \alpha\rho - \beta S ,$$

with $\alpha > 0, \beta \geq 0$.

- Finite-time-blow-up:

$$\lim_{t \rightarrow T} \left(\|\rho(\cdot, t)\|_{L^\infty(\mathbb{R}^n)} + \|S(\cdot, t)\|_{L^\infty(\mathbb{R}^n)} \right) = \infty .$$

Ex: $\rho(\cdot, T) \rightarrow \delta_a$.

Re-scaling

Let us go back to the Othmer-Dunbar-Alt model:

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Re-scaling

$$x' = x/x_0 , \quad t' = t/t_0 , \quad v' = v/v_0 ,$$

$$T' = T/T_0 , \quad S' = S/S_0 , \quad \rho' = \rho/\rho_0 ,$$

$$f' = f/f_0 .$$

Re-scaling

$$\frac{\partial f}{\partial t} + \frac{v_0}{x_0/t_0} v \cdot \nabla f = T_0 v_0^n t_0 \int_V (T f' - T^* f) dv' ,$$

$$\frac{\partial S}{\partial t} = \frac{t_0}{x_0^2} D_0 \Delta S + \frac{\alpha_1 \rho_0 t_0}{S_0} \rho - \alpha_2 t_0 S ,$$

$$\rho = \frac{f_0 v_0^n}{\rho_0} \int_V f dv .$$

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We impose the *diffusive scaling*: $t_0 \approx x_0^2$, normalizations and

$$\varepsilon = \frac{x_0/t_0}{v_0} .$$

Re-scaling

$$\frac{\partial f}{\partial t} + \frac{1}{\varepsilon} v \cdot \nabla f = \frac{1}{\varepsilon^2} \int_V (T_\varepsilon f' - T_\varepsilon^* f) dv' ,$$

$$\frac{\partial S}{\partial t} = \Delta S + \rho - S ,$$

$$\rho = \int_V f dv .$$

The kernel T depends on ε ...

Re-scaling

$$\frac{\partial f_\varepsilon}{\partial t} + \frac{1}{\varepsilon} v \cdot \nabla f_\varepsilon = \frac{1}{\varepsilon^2} \int_V (T_\varepsilon f'_\varepsilon - T_\varepsilon^* f_\varepsilon) dv' ,$$

$$\frac{\partial S_\varepsilon}{\partial t} = \Delta S_\varepsilon + \rho_\varepsilon - S_\varepsilon ,$$

$$\rho_\varepsilon = \int_V f_\varepsilon dv .$$

The solution depends on ε ...

Formal computations

Consider the turning kernel

$$T_\varepsilon[S, \rho] = T_0[S, \rho] + \varepsilon T_1[S, \rho] + \dots$$

such that

$$\begin{aligned} T_0[S, \rho] &= \lambda(S, \rho)(x, t)F(v) , \\ T_1[S, \rho] &= F(v)a(S, \rho)v \cdot \nabla S , \end{aligned}$$

where

$$F > 0 , \quad \int_V F dv = 1 , \quad \int_V v F dv = 0 ,$$

$$\lambda \geq \lambda_{\min} > 0$$

Formal computations

Two possible examples are given by

$$T_\varepsilon[S, \rho] = F(v)\lambda(S, \rho) + \varepsilon F(v)a(S, \rho)v \cdot \nabla S ,$$

$$T_\varepsilon[S, \rho] = \psi(S(x, t), S(x + \varepsilon\mu(\rho)v, t))F(v) ,$$

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We consider the formal expansion of the solutions:

$$f_\varepsilon = f_0 + \varepsilon f_1 + \varepsilon^2 f_2 + \dots ,$$

$$S_\varepsilon = S_0 + \varepsilon S_1 + \varepsilon^2 S_2 + \dots .$$

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We put these expansions in the model, match terms with the same order of ε and solve the resulting system.

Formal computations

In both cases the *formal drift-diffusion limit* is given by the zeroth order equations (Othmer, Hillen)

$$\begin{aligned}\partial_t \rho_0 &= \nabla \cdot (D(S_0, \rho_0) \nabla \rho_0 - \chi(S_0, \rho_0) \rho_0 \nabla S_0) , \\ \partial_t S_0 &= \Delta S_0 + \rho_0 - S_0 ,\end{aligned}$$

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where

$$D(S_0, \rho_0) := \frac{1}{n\lambda(S_0, \rho_0)} \int_V v^2 F dv \mathbb{I} ,$$

$$\chi(S_0, \rho_0) := \frac{a(S, \rho)}{n\lambda(S_0, \rho_0)} \left(\int_V v^2 F dv \right) .$$

General Picture (again...)

	Model $\varepsilon > 0$	Limit model $\varepsilon \rightarrow 0$
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$$\Phi(t) = \lim_{\varepsilon \rightarrow 0} \Phi_\varepsilon(t), \quad t < T$$

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Rigorous results

Theorem (C., Markowich, Perthame, Schmeiser, Hwang, Kang, Stevens, Rodrigues): Consider turning kernels T_ε depending on S_ε , ∇S_ε and ρ_ε under mild assumptions. Then, the solution of the kinetic model $(f_\varepsilon, S_\varepsilon)$ is such that

$$\rho_\varepsilon \rightarrow \rho_0 \text{ in } L^2_{\text{loc}}(\mathbb{R}^n),$$

$$S_\varepsilon \rightarrow S_0 \text{ in } L^q_{\text{loc}}(\mathbb{R}^n), 1 \leq q < \infty,$$

$$\nabla S_\varepsilon \rightarrow \nabla S_0 \text{ in } L^q_{\text{loc}}, 1 \leq q < \infty.$$

where (ρ_0, S_0) is the solution of the associated Keller-Segel model.

Rigorous Results

By *mild assumptions* we mean:

$$\begin{aligned}\phi_\varepsilon^S[\rho, S] &\geq \gamma(1 - \varepsilon\Lambda(\|S\|_{W^{1,\infty}}))FF', \\ \int_V \frac{\phi_\varepsilon^A[\rho, S]^2}{F\phi_\varepsilon^S[\rho, S]} dv' &\leq \varepsilon^2\Lambda(\|S\|_{W^{1,\infty}}),\end{aligned}$$

where

$$\begin{aligned}\phi_\varepsilon^S &:= \frac{T_\varepsilon[S, \rho]F' + T_\varepsilon^*[\rho, S]F}{2}, \\ \phi_\varepsilon^A &:= \frac{T_\varepsilon[S, \rho]F' - T_\varepsilon^*[\rho, S]F}{2},\end{aligned}$$

and other more technical ones.

Global Existence

Theorem (C., Markowich, Perthame, Schmeiser, Hwang, Kang, Stevens): Consider turning kernels such that:

$$\begin{aligned} 0 \leq T_\varepsilon[S_\varepsilon, \nabla S_\varepsilon] &\leq c_1 + c_2 S(x + \varepsilon v, t) + c_3 S(x - \varepsilon v', t) \\ &\quad + c_4 |\nabla S(x + \varepsilon v, t)| + c_5 |\nabla S(x - \varepsilon v, t)|, \\ |\nabla T_\varepsilon[S_\varepsilon, \nabla S_\varepsilon]| &\leq c_2 |\nabla S(x + \varepsilon v, t)| + c_3 |\nabla S(x - \varepsilon v', t)| \\ &\quad + c_4 |\nabla^2 S(x + \varepsilon v, t)| + c_5 |\nabla^2 S(x - \varepsilon v, t)|. \end{aligned}$$

The kinetic model has global existence of solutions.

Global Existence

Let us consider

$$T_\varepsilon[S](x, v, v', t) = \psi(S(x + \varepsilon v, t) - S(x, t))$$

with $\psi \geq \psi_{\min} > 0$, increasing and $\psi(y) \leq Ay + B$.

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with $\psi \geq \psi_{\min} > 0$, increasing and $\psi(y) \leq Ay + B$.
Then the kinetic model has global existence of solution,
converges (in the drift diffusion limit) to the *classical*
Keller-Segel model (which presents blow up).

Global Existence

Theorem (C., Rodrigues): For certain classes of turning kernels $T_\varepsilon[S_\varepsilon, \nabla S_\varepsilon, \rho_\varepsilon]$, such that if $\rho_\varepsilon(x, t) \geq \bar{\rho}$, then

$$T_\varepsilon[S_\varepsilon, \nabla S_\varepsilon, \rho_\varepsilon](x, v, v', t) = T_0[S_\varepsilon, \nabla S_\varepsilon, \rho_\varepsilon](x, v, v', t) ,$$

and for ε small enough, we conclude global existence of solution $(f_\varepsilon, S_\varepsilon)$.

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and for ε small enough, we conclude global existence of solution $(f_\varepsilon, S_\varepsilon)$. Furthermore,

$$\|\rho_\varepsilon(\cdot, t)\|_{L^\infty(\mathbb{R}^n)} \leq \max\{\bar{\rho}, \|\rho^I\|_{L^\infty(\mathbb{R}^n)}\} .$$

Global Existence

Example:

$$T_\varepsilon[S, \rho] = \lambda(S, \rho)F + \varepsilon a(S, \rho)Fv \cdot \nabla S ,$$

$$T_\varepsilon[S, \rho] = \psi(S(x + \varepsilon\mu(\rho)v, t) - S(x, t))F$$

with

$$a(S, \rho) = 0 , \quad \mu(\rho) = 0 , \quad \rho \geq \bar{\rho} > 0 .$$

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Corollary: For the limit Keller-Segel model, we can conclude global existence of solutions. Furthermore, the cell density is bounded. (Hillen, Painter)

$$\partial_t \rho = \nabla \cdot (\nabla \rho - \beta(\rho)\rho \nabla S) , \quad \beta(\rho) = 0 , \quad \rho \geq \bar{\rho} > 0 .$$

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A: *Bad:* Blow up does not exist in nature, we should look for models without blow up.

A: *Good:* Keller-Segel model cannot be valid after certain time, so a realistic model should indicate that it breaks down.

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A: Yes. This was done in the kinetic level. Other possibility is given by Velazquez:

$$\partial_t \rho = \nabla \cdot \left(\nabla \rho - \frac{\rho}{1 + \mu \rho} \nabla S \right)$$

For any $\mu > 0$, there is global existence of solutions. For $\mu = 0$, solutions blow up in finite time.

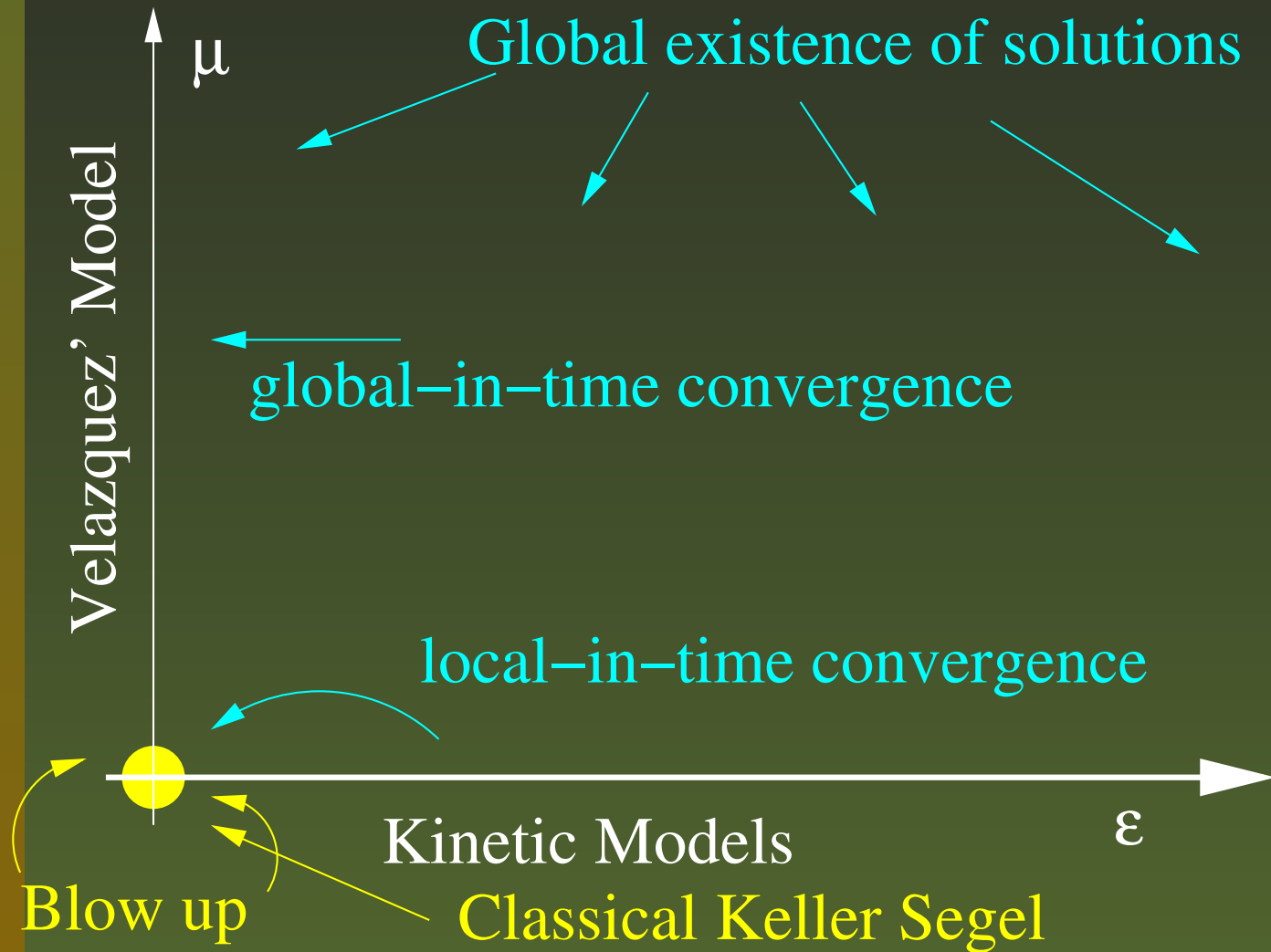
Beyond Keller-Segel

With

$$T_{\varepsilon, \mu}[S, \rho] = \psi \left(S \left(x + \frac{\varepsilon}{1 + \mu\rho} v \right) - S(x, t) \right)$$

the solution exists globally and the drift-diffusion limit (*globally in time*) is the Velazquez' model (C., Kang).

Beyond Keller-Segel



Conclusions

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Closer to *first principles*.

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Detailed description.

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THE END