

# General relative entropy inequality and structured population models

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**1-PBE models** (population balance equation) We study the dynamic of a cell population describes by its density at time  $t$ ,  $n(t, y)$ , in which the intracellular state is characterized by  $y$  (size, age or ...).

## 2-Examples of size structured models

$$\frac{\partial}{\partial t}n(t, y) + \frac{\partial}{\partial y}n(t, y) + B(y)n(t, y) = 4B(2y)n(t, 2y),$$
$$n(t, 0) = 0, \quad n(0, y) = n_0(y),$$

or more generally

$$\frac{\partial}{\partial t}n(t, y) + \frac{\partial}{\partial y}n(t, y) + B(y)n(t, y) = \int_y^\infty b(y, y')n(t, y')dy',$$
$$n(t, 0) = 0, \quad n(0, y) = n_0(y).$$

## 3-Example of age structured model

$$\frac{\partial}{\partial t}n(t, a) + \frac{\partial}{\partial a}n(t, a) + D(y)n(t, y) = 0,$$
$$n(t, 0) = \int B(y)n(t, y)dy, \quad n(0, y) = n_0(y).$$

## I-Aim

Our purpose is to show that in these cases, we have the following asymptotic behaviour such that  $n(t, y) \sim e^{\lambda t} N(y)$  as  $t \rightarrow \infty$  for a

$$\lambda \in \mathbb{R},$$
$$N \in L^1(\mathbb{R}_+), \quad N(y) \geq 0, \quad N \not\equiv 0.$$

### A scattering model :

$$\partial_t f = \int f(y') K(y, y') dy' - \int f(y) K(y', y) dy',$$

then there is mass conservation. Now, if we have the existence of  $N$  solution to  $N(y') K(y, y') = N(y) K(y', y)$ , then for all  $H$  positive convex function

$$\begin{aligned} \partial_t \int H(f/N) N(y) dy &= \int \int [H'(f/N)(y) (\frac{f}{N}(y') - \frac{f}{N}(y)) \\ &\quad + (H(f/N)(y) - H(f/N)(y'))] K(y, y') N(y') dy dy' \leq 0, \end{aligned}$$

(where  $H'(z) = d/dz H(z)$ ) and for some 'good'  $K$  we have  $f \rightarrow N$ .

## Formal conservative law :

Let  $F$  a linear operator (cell division...)

$$\frac{\partial}{\partial t} n(t, y) = F(n).$$

formally we have the following conservative law

$$\frac{\partial}{\partial t} \int n(t, y) \phi(y) dy = \int F(n)(y) \phi(y) dy = \int n(t, y) F^*(\phi)(y) dy = 0$$

as  $F^*(\phi) = 0, \phi > 0$ .

## A cell division model (with constant rate of division):

$$\frac{\partial}{\partial t}n(t, y) + \frac{\partial}{\partial y}n(t, y) + Bn(t, y) = 4Bn(t, 2y),$$

then  $\partial_t \int n(t, y)dy = B \int n(t, y)dy$  and we normalize  $n$  by

$m = ne^{-Bt}$  thus

$$\frac{\partial}{\partial t}m(t, y) = F(m) = -\frac{\partial}{\partial y}m(t, y) - 2Bm(t, y) + 4Bm(t, 2y),$$

$$F^*(\phi) = \frac{\partial}{\partial y}\phi(t, y) - 2B\phi(t, y) + 2B\phi(t, y/2),$$

we have clearly  $F^*(1) = 0$  and  $\int m(t, y)1dy = \int m(0, y)1dy$ .

## II- Entropy

General entropy equation for these non-conservative equations  $\partial_t n = F(n)$  (size or age). Let  $m = ne^{-\lambda t}$  then for all  $H$  convex, positive function, we have

$$\partial_t \int H(m/N) N(y) \phi(y) dy = -D_H(m/N) \leq 0,$$

where  $N, \phi$  are defined by the eigenproblem

$$F(N) = -\lambda N, \quad \int N(y) dy = 1, \quad N(y) \geq 0,$$

$$F^*(\phi) = -\lambda \phi, \quad \int N(y) \phi(y) dy < \infty, \quad \phi > 0,$$

$$\partial_t m = F(m) - \lambda m.$$

## Entropy decreasing term $D_H$ :

For all  $H$  convex, positive function, we have

In the size structured model

$$\begin{aligned} & \partial_t \int H(m/N) N(y) \phi(y) dy = \\ & - \int \int N(y') \phi(y) b(y, y') \left[ H' \left( \frac{m}{N} \right) (y) \left( \frac{m}{N}(y) - \frac{m}{N}(y') \right) + H \left( \frac{m}{N} \right) (y') - H \left( \frac{m}{N} \right) (y) \right] dy' dy. \end{aligned}$$

In the age structured model

$$\begin{aligned} \partial_t \int H(m/N) N(y) \phi(y) dy &= H \left( \int m/N(y) d\mu(y) \right) - \int H(m/N)(y) d\mu(y), \\ d\mu(y) &= b(y) N(y) dy / \int b(y') N(y') dy'. \end{aligned}$$

We only use algebraic calculation and equations satisfied by  $N$ ,  $\phi$ .

### III- Results

Under following assumptions

1- Existence of  $(\lambda, N, \phi)$  solution to the eigenproblem

$$F(N) = -\lambda N, \quad \int N(y)dy = 1, \quad N(y) \geq 0,$$
$$F^*(\phi) = -\lambda\phi, \quad \int N(y)\phi(y)dy < \infty, \quad \phi > 0.$$

2- Some assumptions on  $b(., .)$ ,  $B(.)$  ( $B(.)$ ,  $D(.)$ ) that give  $D_H(f) = 0$  iff  $f$  is constant,

we have

$$n(t, .)e^{-\lambda t} \rightarrow CN(y) \quad \text{in } L^p(\mathbb{R}_+, N(y)\phi(y)dy).$$

Moreover we have  $\int n(t, y)e^{-\lambda t}\phi(y)dy = \int n(0, y)\phi(y)dy$ .



## IV- Link with evolution

We obtain  $n(t, y) \sim e^{\lambda t} N(y)$ , thus the  $\lambda$  is a measure of adaptation. For example let

$n_1(t, y)$  solution to the size struct. with  $b(y, y') = \delta_{y=y'/2} + \delta_{y=y'/2}$ ,

$n_2(t, y)$  solution to the size struct. with  $b(y, y') = \delta_{y=y'/3} + \delta_{y=2y'/3}$ ,

then  $n_1 \sim N_1 e^{\lambda_1 t}$  and  $n_2 \sim N_2 e^{\lambda_2 t}$ . In this case we have  $\lambda_1 = \lambda_2 = 1$  and the two species evolve with the same exponential growth.

In general, the exact value of  $\lambda$  is impossible to find and the link between parameters  $(b, B)$  and  $\lambda$  is not clear. Nevertheless one can find numerically the solution of the eigenproblem.

# V- Summary

## 1-Advantage

**a-Entropy equation** We can compute  $\partial_t \int N \phi H(n/N) dy$  independently of the existence of  $N, \phi$ -**algebraic calculation**.

**b-Existence**  $N, \phi, \lambda$  In discrete and compact cases we have the existence of the solution to the eigenproblem.

**c-Result**  $n \sim e^{\lambda t} N(y)$  .

## 2-Disadvantage

**a-Speed rate (for convergence)** Hidden in  $D_H(\cdot)$ ,

**b- $\lambda$**  No accurate information in general.

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