

# Optimal control for a conservation law modeling the development of ovulation

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# Outline

- Introduction of the model and related control problems
- Some numerical results
- Theoretical result for Dirac masses based on Pontryagin's Maximum Principle (PMP)
- Numerical results based on PMP
- Optimal control result for PDE case
- Open problems
- References

The cell population in a follicle is represented by cell density functions  $\rho_{j,k}(t, x, y)$  defined on each cellular phase  $Q_{j,k}$  with age  $x$  and maturity  $y$ , which satisfy the following conservation laws

$$\frac{\partial \rho_{j,k}(t, x, y)}{\partial t} + \frac{\partial \rho_{j,k}(t, x, y)}{\partial x} + \frac{\partial (h(y, u) \rho_{j,k}(t, x, y))}{\partial y} = 0, \quad \text{in } Q_{j,k} \quad (1.1)$$

Here  $k = 1, \dots, N$ , and  $N$  is the number of consecutive cell cycles.  $j = 1, 2, 3$  denotes Phase 1, Phase 2 and Phase 3.

$$h(y, u) = -y^2 + (c_1 y + c_2) u, \quad (1.2)$$

with  $c_1$  and  $c_2$  given positive constants.

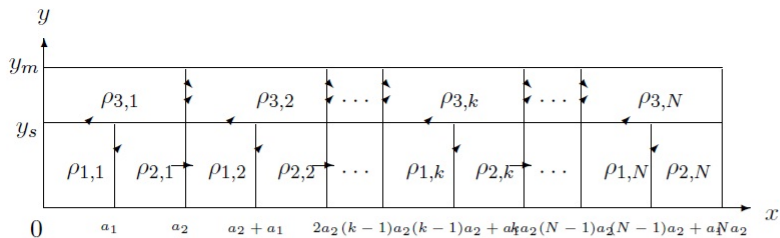


Figure 1:



## related control problems

- We define

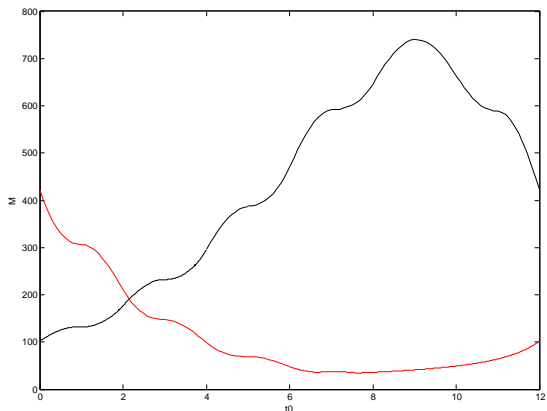
$$M(t) := \sum_{j=1}^3 \sum_{k=1}^N \int_0^{+\infty} \int_0^{+\infty} y \rho_{j,k}(t, x, y) dx dy \quad (1.3)$$

as the follicular maturity.

- The control  $u(M(t))$ .
- Ovulation is triggered when the maturity reaches a given threshold value  $M_s$ . Hence, the optimal control problem is, for fixed observed time  $t_1$  to maximize the maturity  $M(t_1)$ .
- Proliferative cells leave the cycle in an irreversible way, we get the restraint of control  $u \in [w, 1]$  with  $w \in (0, 1)$ .

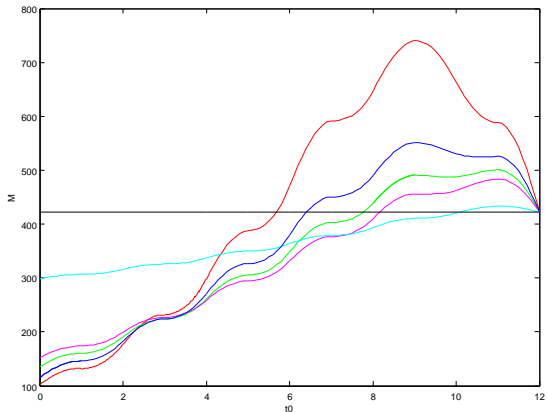
## switching direction

black :  $u = w \rightarrow u = 1$ ; red:  $u = 1 \rightarrow u = w$



# Numerical result of optimal bang-bang control

red:  $u = w \rightarrow u = 1$ , black:  $u = w$



## simplified model

Consider the following balance law

$$\rho_t + \rho_x + (h(y, u)\rho)_y = c_s \chi(y), \quad t \geq 0, x \geq 0, y \geq 0, \quad (3.1)$$

with  $c_s$  a given positive constant, we denote

$$a(y) = -y^2, \quad b(y) = c_1 y + c_2. \quad (3.2)$$

$$h(y, u) = a(y) + b(y)u. \quad (3.3)$$

$\chi(y)$  is a characteristic function

$$\chi(y) = \begin{cases} 1, & \text{if } y \in [0, y_s), \\ 0, & \text{if } y \in (y_s, \infty), \end{cases} \quad (3.4)$$

**(BC)**

$$\rho(t, 0, y) = \rho(t, x, 0) = 0, \quad \forall x \geq 0, y \geq 0. \quad (3.5)$$

**(IC)**

The initial condition  $\rho_0(x, y)$  is given as a positive Borel measure with compact support  $\subset [0, 1]^2$ .

For any admissible control  $u \in L^\infty([t_0, t_1]; [w, 1])$ , the cost function is

$$J(u) = - \iint_{[0, +\infty) \times [0, +\infty)} y d\rho(t_1, x, y). \quad (3.6)$$

## one Dirac mass

We consider the following optimal control problem ( $\mathcal{P}$ ):

$$\begin{cases} \dot{x} = f(x, u), & u \in L^\infty([t_0, t_1]; [w, 1]), & t \in [t_0, t_1], \\ x(t_0) = x^0, \\ J(u) = \int_{t_0}^{t_1} (p(x, u) + q(x) \chi(x_2)) dt \end{cases} \quad (3.7)$$

where

$$p(x, u) := -(a(x_2) + b(x_2) u) x_3, \quad q(x) := -c_s x_2 x_3. \quad (3.8)$$

$$f = \begin{pmatrix} 1 \\ a(x_2) + b(x_2) u \\ c_s \chi(x_2) x_3 \end{pmatrix}$$

$x = (x_1, x_2, x_3)^{tr} \in R^3$ ,  $x_1$  denotes the age,  $x_2$  denotes the maturity and  $x_3$  denotes the mass.

The main difficulty of problem  $(\mathcal{P})$  is that both  $f$  and the integrand  $\chi$  of the functional  $J$  are discontinuous. First, we proved the existence of optimal control to problem  $(\mathcal{P})$  by approximating method.

### Theorem

*The infimum of the functional  $J$  in  $L^\infty([t_0, t_1]; [w, 1])$  is achieved, i.e., there exists  $u \in L^\infty([t_0, t_1]; [w, 1])$  such that*

$$J(u) = \inf_{u \in L^\infty([t_0, t_1]; [w, 1])} J(u).$$

## Theorem

For any measurable optimal control  $u_*$  to problem  $(\mathcal{P})$ , the following property holds:

There exists  $t' \in [t_0, t_1)$  such that

$$u_* = w \text{ in } (t_0, t') \text{ and } u_* = 1 \text{ in } (t', t_1). \quad (3.9)$$

Furthermore, under the assumption that

$$2y_s - c_1 > 0 \quad \text{and} \quad c_s > \frac{a(y_s) + b(y_s)}{y_s}, \quad (3.10)$$

this optimal switch time  $t'$  is the exit time  $\hat{t}$ .



# Necessary optimal conditions for optimal control (PMP)

Necessary optimal conditions (A. I. Smirnov, 2008). Let us define the Hamilton-Pontryagin function and the Hamiltonian as

$$\mathcal{H}(x, u, \psi, \psi^0) := \langle f(x, u), \psi \rangle + \psi^0 \left( p(x, u) + q(x)\chi(x_2) \right), \quad (3.11)$$

$$H(x, \psi, \psi^0) = \max_{u \in U} \mathcal{H}(x, u, \psi, \psi^0). \quad (3.12)$$

Let us denote  $\hat{t}$  as  $x_{*2}(\hat{t}) = y_s$ .

Let  $u_* \in L^\infty([t_0, t_1]; [w, 1])$  and  $x_* = (x_{*1}, x_{*2}, x_{*3})^{tr}$  be the optimal control and the corresponding optimal trajectory in problem  $(\mathcal{P})$ . Then there exist a function  $\psi = (\psi_1, \psi_2, \psi_3)^{tr}$ ,  $\psi_1 \in W^{1,\infty}(t_0, t_1)$ ,  $\psi_2 \in W^{1,\infty}(t_0, \hat{t}) \cup (\hat{t}, t_1)$ , and  $\psi_3 \in W^{1,\infty}(t_0, t_1)$  such that the following conditions hold:

(a) The function  $\psi$  is a solution to the adjoint system:

$$\dot{\psi}_1(t) = 0, \quad (3.13)$$

$$\begin{aligned} \dot{\psi}_2(t) = & - \left( \frac{\partial a(x_{*2}(t))}{\partial x_2} + \frac{\partial b(x_{*2}(t))}{\partial x_2} u_*(t) \right) \psi_2(t) \\ & + \frac{\partial p(x_*(t), u_*(t))}{\partial x_2} + \frac{\partial q(x_*(t))}{\partial x_2} \chi(x_{*2}(t)), \quad t \neq \hat{t}, \end{aligned} \quad (3.14)$$

$$\begin{aligned} \dot{\psi}_3(t) = & -c_s \chi(x_{*2}(t)) \psi_3(t) + (a(x_{*2}(t)) + b(x_{*2}(t))u_*(t)) \\ & + c_s x_{*2}(t) \chi(x_{*2}(t)). \end{aligned} \quad (3.15)$$

(b) The jump when the  $x_{*2} = y_s$ :

$$\psi_2(\hat{t} + 0) - \psi_2(\hat{t} - 0) \in c_s x_3(\hat{t}) \left( \frac{y_s + \psi_3(\hat{t})}{a(y_s) + b(y_s)}, \frac{y_s + \psi_3(\hat{t})}{a(y_s) + b(y_s)w} \right).$$

$$\psi_1(t_1) = \psi_2(t_1) = \psi_3(t_1) = 0.$$

(c) The maximum condition holds:

$$H(x_*(t), \psi) \stackrel{a.e.}{=} \mathcal{H}(x_*(t), u_*(t), \psi).$$

# proof of the Theorem

Now, the Hamiltonian becomes

$$H(t) = (a(x_2) + c_s \chi x_2) x_3 + \psi_1 + a(x_2) \psi_2 + c_s \chi x_3 \psi_3 + b(x_2)(x_3 + \psi_2) u.$$

Noting that  $b(x_2) > 0$ , one has

$$u_*(t) = 1 \quad \text{if} \quad x_3(t) + \psi_2(t) > 0, \quad (3.16)$$

$$u_*(t) = w \quad \text{if} \quad x_3(t) + \psi_2(t) < 0. \quad (3.17)$$

Under the assumption that  $2y_s - c_1 > 0$ , we have

$$(x_3 + \psi_2)(\hat{t} - 0) \leq x_3(t_1) \left(1 - c_s \frac{y_s}{a(y_s) + b(y_s)}\right). \quad (3.18)$$

Under the assumption that  $c_s > \frac{a(y_s) + b(y_s)}{y_s}$ , we have

$$(x_3 + \psi_2)(\hat{t} - 0) < 0. \quad (3.19)$$

Hence,

$$(x_3 + \psi_2)(t) < 0, \quad t \in [t_0, \hat{t}). \quad (3.20)$$

Moreover,

$$(x_3 + \psi_2)(t) > 0, \quad t \in (\hat{t}, t_1]. \quad (3.21)$$

Above all, we have proved the Theorem.

# proof of the necessary optimal conditions

**Step 1:** Mollifier the characteristic  $\chi$ .

**Step 2:** Let  $u_*$ ,  $x_*$  be an optimal pair in problem  $(\mathcal{P})$ . Take a sequence  $\{z_i\}$ ,  $i = 1, 2, \dots$ , of functions  $z_i \in C^1[t_0, t_1]$  that satisfy the following conditions

$$z_i \rightarrow u_* \text{ in } L^2[t_0, t_1] \text{ as } i \rightarrow \infty, \quad (3.22)$$

$$\sup_{t_0 \leq t \leq t_1} \|z_i(t)\| \leq \|U\| + 1, \quad i = 1, 2, \dots, \quad (3.23)$$

$$\sup_{t_0 \leq t \leq t_1} \|\dot{z}_i(t)\| \leq \sigma_i < \infty. \quad (3.24)$$

We may assume without loss of generality that  $\sigma_i \rightarrow \infty$  as  $i \rightarrow \infty$ .

**Step 3:** Now consider the following sequence of auxiliary optimal control problems  $(\mathcal{P}_i)$

$$\begin{cases} \dot{x} = f_i(x, u), & u \in L^\infty([t_0, t_1]; [w, 1]), & t \in [t_0, t_1], \\ x(t_0) = x^0, \\ J_i(u) = \int_{t_0}^{t_1} \left( p(x, u) + q(x) \chi_i(x_2) \right) dt + \frac{1}{1 + \sigma_i} \int_{t_0}^{t_1} \|u(t) - z_i(t)\|^2 dt. \end{cases} \quad (3.25)$$

Here

$$f_i = \begin{pmatrix} 1 \\ a(x_2) + b(x_2) u \\ c_s \chi_i(x_2) x_3 \end{pmatrix}$$

For any  $i = 1, 2, \dots$ , problem  $(\mathcal{P}_i)$  is a smooth optimal control problem. Hence, there exists an optimal control  $u_i$  in problem  $(\mathcal{P}_i)$  (L. Cesari, 1983). Let  $x_i$  be the corresponding optimal trajectory. We have the following result

### Lemma

*The following relations hold as  $i \rightarrow \infty$*

$$u_i \rightarrow u_* \quad \text{in } L^2[t_0, t_1], \quad (3.26)$$

$$x_i \rightarrow x_* \quad \text{in } C^0[t_0, t_1]. \quad (3.27)$$



Suppose that  $x_i$  and  $u_i$  is an optimal pair in problem  $(\mathcal{P}_i)$ . Define the Hamilton-Pontryagin function and the Hamiltonian for problem  $(\mathcal{P}_i)$  as follows

$$\begin{aligned} \mathcal{H}_i(t, x, u, \psi, \psi^0) = & \langle f_i(x, u), \psi \rangle + \psi^0(p(x, u) + q(x)\chi_i(x_2)) \\ & + \psi^0\left(\frac{1}{1 + \sigma_i} \|u(t) - z_i(t)\|^2\right), \end{aligned}$$

and

$$H_i(t, x, \psi, \psi^0) = \max_{u \in U} \mathcal{H}_i(t, x, u, \psi, \psi^0).$$

By Pontryagin's maximum principle, there exists a number  $\psi_i^0 \leq 0$  and an absolutely continuous function  $\psi_i$  on  $[t_0, t_1]$  such that

$$\begin{aligned} \dot{\psi}_i(t) &\stackrel{a.e.}{=} - \left[ \frac{\partial f_i(x_i(t), u_i(t))}{\partial x} \right]^* \psi_i - \psi_i^0 \frac{\partial p(x_i(t), u_i(t))}{\partial x} \\ &\quad - \psi_i^0 \left( \frac{\partial q(x_i(t))}{\partial x} \chi_i(x_i(t)) + q(x_i(t)) \frac{\partial \chi_i(x_i(t))}{\partial x} \right), \\ \psi_i(t_1) &= 0, \end{aligned} \tag{3.28}$$

and

$$H_i(t, x_i(t), \psi_i(t), \psi_i^0) \stackrel{a.e.}{=} \mathcal{H}_i(t, x_i(t), u_i(t), \psi_i(t), \psi_i^0). \tag{3.29}$$

Passing to the limit  $i \rightarrow \infty$  in necessary optimal conditions for problem  $(\mathcal{P}_i)$ , finally we prove the necessary optimal conditions for problem  $(\mathcal{P})$ .

## n Dirac masses

Using the same method as one Dirac mass, we get similar result that

### Theorem

*Under the assumption that*

$$2y_s - c_1 > 0 \quad \text{and} \quad c_s > \frac{a(y_s) + b(y_s)}{y_s}, \quad (3.30)$$

*For any optimal control  $u_*$ , the following property holds*

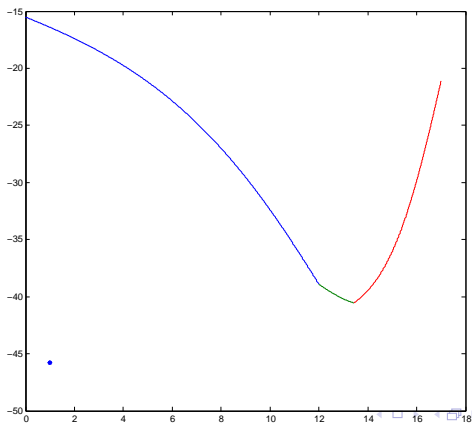
*There exists  $t' \in (t_0, t_1)$  such that*

$$u_* = w \text{ in } (t_0, t') \text{ and } u_* = 1 \text{ in } (t', t_1). \quad (3.31)$$

## counter example with small $c_s$

### Remark

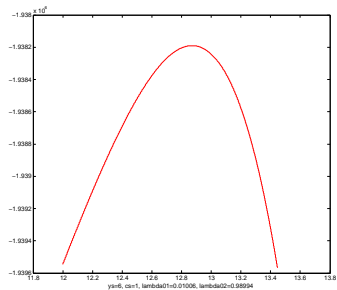
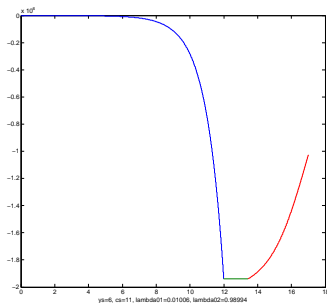
The assumption (3.30) is important to guarantee that the optimal switch time is once.



# nonuniqueness of the optimal control for two Dirac masses

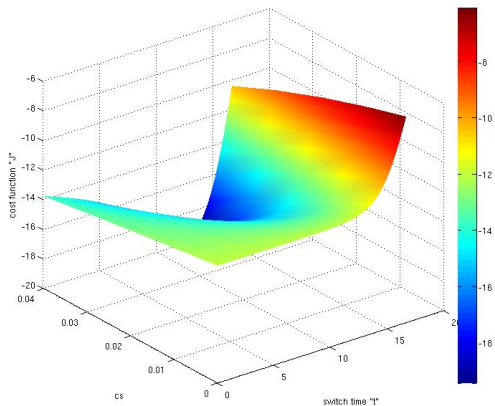
## Remark

For more than one Dirac masses, the optimal control is not unique.



optimal control for one Dirac mass with different  $c_s$ 

For one Dirac mass, when  $c_s$  is small, the optimal control is always  $u = 1$ ; when  $c_s$  is large, the optimal control is  $u = w \rightarrow u = 1$ .



Recall the PDE case

$$\rho_t + \rho_x + (h(y, u)\rho)_y = c_s \chi(y), \quad t \geq 0, x \geq 0, y \geq 0. \quad (5.1)$$

The cost function is

$$J(u) = - \iint_{[0, +\infty) \times [0, +\infty)} y d\rho(t_1, x, y). \quad (5.2)$$



We have the following result

### Theorem

*Under the assumption that*

$$2y_s - c_1 > 0 \quad \text{and} \quad c_s > \frac{a(y_s) + b(y_s)}{y_s}, \quad (5.3)$$

*we have that among all admissible controls  $u \in L^\infty([t_0, t_1]; [w, 1])$ , there exists an optimal control  $u_*$  to (5.2) such that the following property holds*

*There exists  $t' \in (t_0, t_1)$  such that*

$$u_* = w \text{ in } (t_0, t') \text{ and } u_* = 1 \text{ in } (t', t_1). \quad (5.4)$$

**Step 1:** There exists a sequence

$$\rho_0^n = \sum_{i=1}^n \lambda_0^i \delta_{x_0^i, y_0^i}, \quad (5.5)$$

such that for any given  $\varphi \in C^0(K)$  we have

$$(\rho_0^n - \rho_0) \varphi \rightarrow 0 \quad \text{as } n \rightarrow \infty. \quad (5.6)$$

The cost function is

$$J(\rho_0^n, u) = - \sum_{i=1}^n y_i(t_1, u) \lambda_i(t_1, u). \quad (5.7)$$

For any  $u \in L^\infty([t_0, t_1]; [w, 1])$ , it is easy to prove that

$$\lim_{n \rightarrow \infty} J(\rho_0^n, u) = J(\rho_0, u). \quad (5.8)$$

**Step 2:** We assume that for each  $\rho_0^n$ , there exists an optimal control  $u_*^n$  such that

$$u_*^n := w \text{ in } (t_0, t'_n), \quad u_*^n := 1 \text{ in } (t'_n, t_1). \quad (5.9)$$

Without loss of generality, we may assume there exists  $t' \in [t_0, t_1]$  such that

$$t'_n \rightarrow t' \text{ as } n \rightarrow \infty. \quad (5.10)$$

Let  $u_*$  be defined as

$$u_* := w \text{ in } (t_0, t'), \quad u_* := 1 \text{ in } (t', t_1). \quad (5.11)$$

Then we prove that






$$\lim_{n \rightarrow \infty} J(\rho_0^n, u_*^n) = J(\rho_0, u_*). \quad (5.12)$$






Combining (5.8) and (5.12), we have proved that  $u_*$  defined an optimal control.

# open problems

- For PDE case, can we get the result that each measurable optimal control is bang-bang control?
- For the moment, we consider the open loop problem, what about the close loop problem, e.x.  $u(t) = u(t, M^1(t))$ ?

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**Thanks for your attention!**