

# Global asymptotic stability of the periodic solution for a periodic model of hematopoiesis with impulses

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# Delay differential equations

- For  $\tau \in \mathbb{R}^+$  and  $m \in \mathbb{N}$ , consider

$$\mathcal{C} := C([- \tau, 0]; \mathbb{R}^m) = \left\{ \varphi : [- \tau, 0] \rightarrow \mathbb{R}^m \mid \varphi \text{ is continuous} \right\}$$

with the norm

$$\|\varphi\| = \sup_{\theta \in [- \tau, 0]} |\varphi(\theta)|_{\mathbb{R}^m}.$$

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- The real space  $(\mathcal{C}, \|\cdot\|)$  is a Banach space.

- ▶ For  $D \subseteq \mathbb{R} \times \mathcal{C}$  and  $f : D \rightarrow \mathbb{R}^m$  continuous, we call a delay differential equation to the equation

$$x'(t) = f(t, x_t).$$

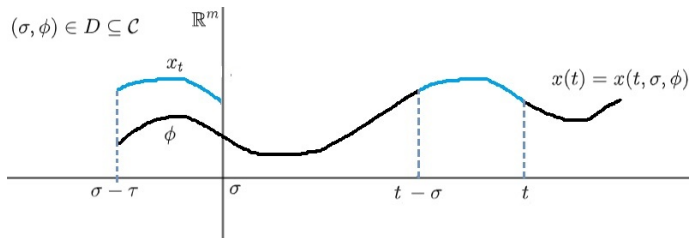
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- Let  $\sigma \in \mathbb{R}$ ,  $b \in (\sigma, +\infty]$ , and  $x : [\sigma - \tau, b] \rightarrow \mathbb{R}^m$  a continuous function.

For  $t \in [\sigma, b]$ , the function  $x_t \in \mathcal{C}$  is defined by

$$x_t(\theta) = x(t + \theta), \quad \forall \theta \in [-\tau, 0].$$



For  $(\sigma, \phi) \in D$ , consider the initial value problem

$$\begin{cases} x'(t) = f(t, x_t) \\ x_\sigma = \phi \end{cases} \quad (1)$$

- As  $f$  is continuous, then IVP (1) has a solution.

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- ▶ If  $x(t)$  is a maximal solution of (1), then, for each compact  $W \subseteq D$ , there is  $t_W \in \mathbb{R}$  such that  $(t, t_W) \notin D$  for  $t \geq t_W$ .



# Biological models

In what follows, we only consider the scalar case ( $m = 1$ ) and non-negative time ( $t \geq 0$ ).

- Scalar biological models

$$x'(t) = -a(t)x(t) + f(t, x_t), \quad t \geq 0,$$

where:

- $a : [0, \infty) \rightarrow \mathbb{R}^+$  is a periodic functions;
- $\forall \phi \in \mathcal{C}, t \mapsto f(t, \phi)$  is a periodic functions.
- Mortality:  $a(t)x(t)$
- Birth:  $f(t, x_t)$

# Impulsive biological models

- Scalar impulsive delay differential equation

$$\begin{cases} x'(t) = -a(t)x(t) + f(t, x_t), & 0 \leq t \neq t_k, \\ x(t_k^+) = x(t_k) + I_k(x(t_k)), & k = 1, 2, \dots \end{cases}, \quad (2)$$

where

- $(t_k)_{k \in \mathbb{N}}$  such that  $0 < t_k \nearrow +\infty$ ;
- $I_k : \mathbb{R} \rightarrow \mathbb{R}$  continuous;
- $a : [0, \infty) \rightarrow (0, \infty)$  continuous;
- $f : [0, \infty) \times PC \rightarrow [0, \infty)$  with some regularities.

# Impulsive biological models

- Scalar impulsive delay differential equation

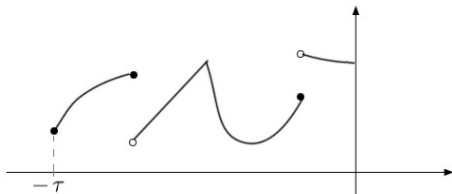
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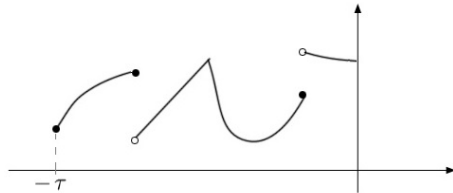
- $(t_k)_{k \in \mathbb{N}}$  such that  $0 < t_k \nearrow +\infty$ ;
  - $I_k : \mathbb{R} \rightarrow \mathbb{R}$  continuous;
  - $a : [0, \infty) \rightarrow (0, \infty)$  continuous;
  - $f : [0, \infty) \times PC \rightarrow [0, \infty)$  with some regularities.
- Note that:

$$x(t_k^+) - x(t_k) = I_k(x(t_k)), \quad \forall k \in \mathbb{N}.$$

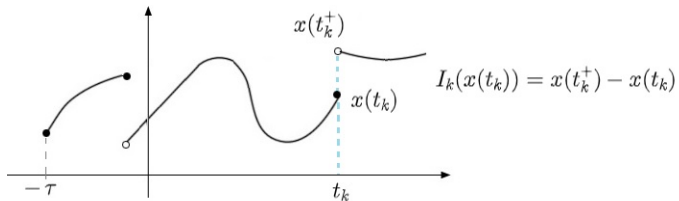
►  $PC = \left\{ \varphi : [-\tau, 0] \rightarrow \mathbb{R} \mid \varphi \text{ is piecewise continuous} \right\}$



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► For  $x(t)$  a solution of (2)



The key step in the study of stability of many impulsive models. For  $x(t)$  a solution of (2) on  $[0, \infty)$ , define

$$\begin{aligned} y(t) &= \prod_{k:0 \leq t_k < t} \frac{x(t_k)}{x(t_k) + I_k(x(t_k))} x(t) \\ &= \prod_{k:0 \leq t_k < t} J_k(x(t_k)) x(t), \end{aligned}$$

where  $J_k(u) = \frac{u}{u + I_k(u)}$ ,  $\forall k \in \mathbb{N}$ ,  $u \neq 0$ .

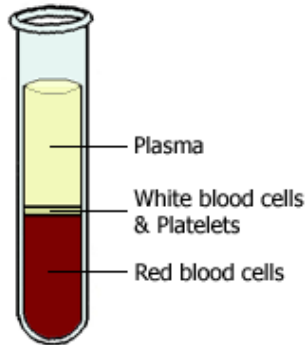
The function  $y(t)$  is continuous and it is solution of

$$y'(t) = -a(t)y(t) + \prod_{k:0 \leq t_k < t} J_k(x(t_k)) f(t, x_t), \quad 0 \leq t \neq t_k.$$

# Processo hematopoiese

Processo de produção, multiplicação e especialização das células do sangue na medula óssea.

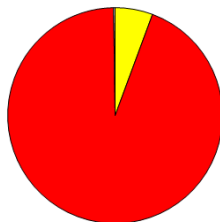
# Constituintes do sangue



O sangue é composto por 55 % plasma e 45% células sanguíneas.



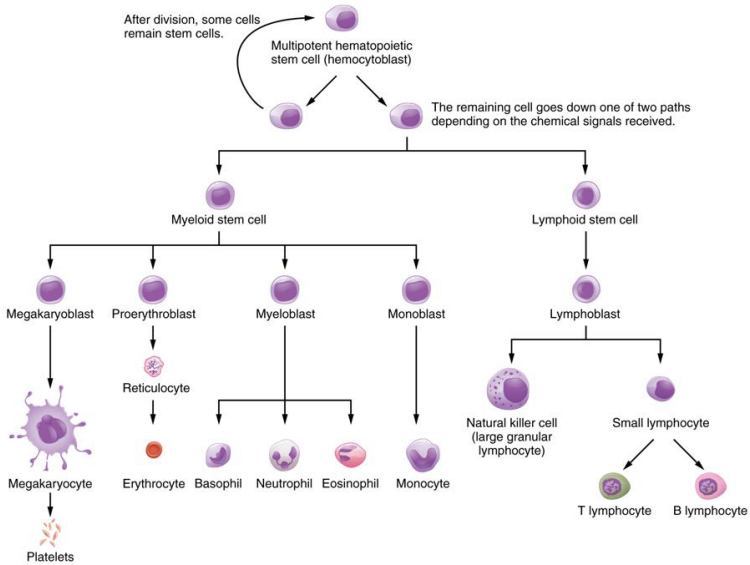
Tipo de células	células/ $1\mu l$ [2]
thrombocytes (Plaquetas)	$15 \times 10^4 - 40 \times 10^4$
erythrocytes (Glob. vermelhos)	homem $43 \times 10^5 - 59 \times 10^5$ mulher $35 \times 10^5 - 55 \times 10^5$
leukocytes (Glob. brancos)	4500 - 11000



- Plaquetas: 5,6%
- Glob. Vermelhos: 94,1%
- Glob. Brancos: 0,3%

$$1\mu l = 1mm^3$$

[2] L. Dean, *Blood Group and Red Cell Antigens*, National Center for Biotechnology Information (US), 2005.



## Tempo de gestação na medula óssea

Tipo de células	Tempo de gestação
thrombocytes (Plaquetas)	$\simeq 7$ dias [3]
erythrocytes (Glob. vermelhos)	$\simeq 6$ dias [4]
neutrophils (60% dos Glob. brancos)	$\simeq 15$ dias [5]

- [3] G.P. Langlois, M. Craig, A.R. Humphries et al., *Normal and pathological dynamics of platelets in humans*, J. Math. Biol. 75 (2017), 1411–1462.
- [4] J. Bélair, M. C. Mackey, J. M. Mahaffy, *Age-structured and two-delay models for erythropoiesis*, Math. Biosci. 128 (1995), 317–346.
- [5] Y. Yan, J. Sugie, *Existence regions of positive periodic solutions for a discrete hematopoiesis model with unimodal production functions* Appl. Math. Model. 68 (2019), 152–168.

# Hematopoiesis models

Mackey and Glass [1], proposed the following models to describe the hematopoiesis process (the process of production, multiplication, and specialization of blood cells in the bone marrow):

- ▶ Hematopoieses with monotone production rate

$$z'(t) = -\gamma z(t) + \frac{F_0 \eta^n}{\eta^n + z(t - \tau)^n}, \quad n > 0; \quad (3)$$

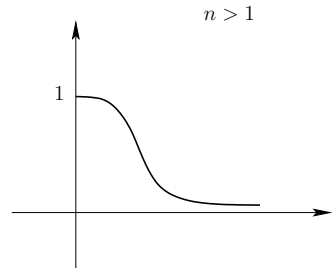
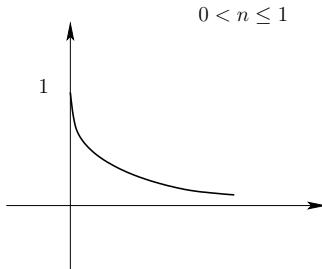
- ▶ Hematopoiesis with unimodal production rate

$$z'(t) = -\gamma z(t) + \frac{F_0 \eta^n z(t - \tau)}{\eta^n + z(t - \tau)^n}, \quad n > 1; \quad (4)$$

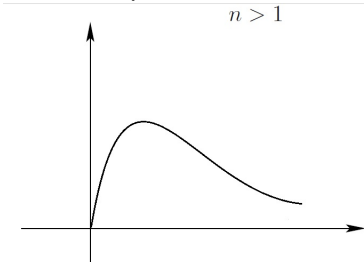
$z(t)$  density of cells at time  $t$ ;  $\tau$  time delay;  $\gamma$  destruction rate;  
 $F_0$  maximal production rate (only for (3));  $\eta$  a shape parameter.

[1] M.C.Mackey, L. Glass, Science 197 (1977) 287-289.

► Monotone production rate



► Unimodal production rate



# Hematopoiesis model with several delays

$$y'(t) = -a(t)y(t) + \sum_{i=1}^m \frac{\beta_i(t)}{1 + y(t - \tau_i(t))^n}, \quad t \geq 0$$

**Notation:**  $\tau(t) = \max_i \tau_i(t)$  and  $\bar{\tau} = \sup_t \tau(t)$

# Hematopoiesis model with linear impulses

For  $(t_k)_k$  an increasing sequence such that  $t_k \rightarrow \infty$ , we consider

$$\begin{cases} y'(t) = -a(t)y(t) + \sum_{i=1}^m \frac{\beta_i(t)}{1 + y(t - \tau_i(t))^n}, & 0 \leq t \neq t_k, \\ y(t_k^+) = (1 + b_k)y(t_k), & k \in \mathbb{N} \end{cases} \quad (5)$$

The impulsive function is linear:  $I_k(u) = b_k u$ . In fact,

$$y(t_k^+) = (1 + b_k)y(t_k) \Leftrightarrow y(t_k^+) = y(t_k) + b_k y(t_k)$$

$$PC_0^+ = \left\{ \varphi \in PC : \varphi(\theta) \geq 0 \text{ for } \theta \in [-\bar{\tau}, 0), \varphi(0) > 0 \right\}$$

► **Periodic Hematopoiesis model with linear impulses**

$$\begin{cases} y'(t) = -a(t)y(t) + \sum_{i=1}^m \frac{\beta_i(t)}{1 + y(t - \tau_i(t))^n}, & 0 \leq t \neq t_k, \\ y(t_k^+) = (1 + b_k)y(t_k), & k \in \mathbb{N} \end{cases}$$



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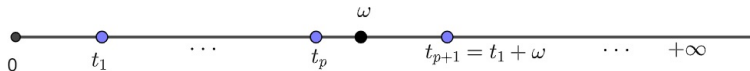
- **(H1)**  $a, \beta_i \in C(\mathbb{R}; (0, \infty))$  and  $\tau_i \in C(\mathbb{R}; [0, \infty))$  are  $\omega$ -periodic, for some  $\omega > 0$ ;

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- **(H2)**  $\exists p \in \mathbb{N}$  such that  $0 < t_1 < \dots < t_p < \omega$  and

$$t_{k+p} = t_k + \omega, \quad b_{k+p} = b_k, \quad k \in \mathbb{N};$$



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- **(H3)**  $1 + b_k > 0, \forall k \in \mathbb{N}$ ;
- **(H4)**  $\prod_{k=1}^p (1 + b_k) < e^{\int_0^\omega a(t)dt}$

# Goal

To establish sufficient conditions for global asymptotic stability (GAS) of a positive  $\omega$ -periodic solution of (5).

# Existence of periodic solution

- **Theorem 2** Faria & Oliveira [3]:  
Assume **(H1)-(H4)**.  
Then system (5) has at least one positive  $\omega$ -periodic solution.

[3] T. Faria and J.J. Oliveira, *Existence of positive periodic solution for scalar delay differential equations with and without impulses*, J. Dyn. Differ. Equ., 31 (2019), 1223-1245.

# Existence of periodic solution

- ▶ **Theorem 2** Faria & Oliveira [3]:  
Assume **(H1)-(H4)**.  
Then system (5) has at least one positive  $\omega$ -periodic solution.
- ▶ In what follows, we fix  $y^*(t)$  a positive  $\omega$ -periodic solution of system (5).

[3] T. Faria and J.J. Oliveira, *Existence of positive periodic solution for scalar delay differential equations with and without impulses*, J. Dyn. Differ. Equ., 31 (2019), 1223-1245.

# Global asymptotic stability

In [4,5], global stability criteria were obtained for the general impulsive model

$$\begin{cases} x'(t) = -a(t)x(t) + \sum_{i=1}^m f_i(t, x(t - \tau_i(t))), & t \neq t_k, \\ x(t_k^+) = (1 + b_k)x(t_k), & k \in \mathbb{N} \end{cases}, \quad (6)$$

where, for each  $i$ ,  $f_i : [0, \infty) \times \mathbb{R} \rightarrow \mathbb{R}$  is piecewise continuous and continuous in the second variable.

- ▶ [4] T. Faria and J.J. Oliveira, *On stability for impulsive delay differential equations and applications to a periodic Lasota-Ważewska model*, Disc. Cont. Dyn. Systems Series B, 21 (2016), 2451-2472.
- ▶ [5] T. Faria and J.J. Oliveira, *A note on stability of impulsive scalar delay differential equations*, Electron. J. Qual. Theory Differ. Equ., Paper No. 69 (2016), 1-14.



- ▶ **Theorem 5** Faria & Oliveira, [4,5]:  
Assume **(H2)-(H3)** and  $a(t) \not\equiv 0$   $\omega$ -periodic continuous.  
The zero solution of (6) is globally asymptotically stable if

(A1) (**Yorke Condition**)

(A2) ( $\frac{3}{2}$ -**Condition**)

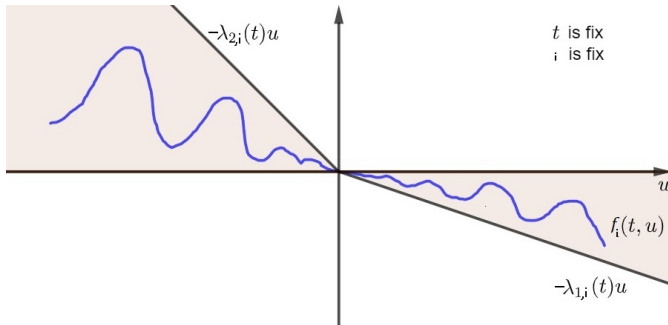
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► **Theorem 5** Faria & Oliveira, [4,5]:

Assume **(H2)**-**(H3)** and  $a(t) \not\equiv 0$   $\omega$ -periodic continuous.  
The zero solution of (6) is GAS if

(A1) (**Yorke Condition**) There are  $\lambda_{1,i}, \lambda_{2,i} : [0, \infty) \rightarrow [0, \infty)$  piecewise continuous such that, for  $t \geq 0$  and  $u \in \mathbb{R}$ ,

$$-\lambda_{1,i}(t) \max\{u, 0\} \leq f_i(t, u) \leq \lambda_{2,i}(t) \max\{-u, 0\};$$



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(A1) (**Yorke Condition**)

(A2) ( $\frac{3}{2}$ -**Condition**) There is  $T > 0$  such that

$$\alpha_1^* \alpha_2^* < 1 \text{ or } \alpha_1 \alpha_2 < \frac{9}{4} \quad (7)$$

where  $\alpha_j^* = \sup_{t \geq T} \alpha_j^*(t)$ ,  $\alpha_j = \sup_{t \geq T} \alpha_j^*(t) e^{\int_{t-\tau(t)}^t a(u) du}$  ( $j = 1, 2$ ),

$$\alpha_j^*(t) = \int_{t-\tau(t)}^t \sum_{i=1}^m \lambda_{j,i}(s) B_i(s) e^{-\int_s^t a(u) du} ds, \quad j = 1, 2.$$

$$\text{with } B_i(s) = \prod_{k: t-\tau_i(t) \leq t_k < t} (1 + b_k)^{-1}, \quad i = 1, \dots, m.$$

## Proof of the main results (idea)

- We translate the positive  $\omega$ -periodic solution of (5),  $y^*(t)$ , to the origin with the change  $x(t) = y(t) - y^*(t)$ .

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- ▶ System (5) is transformed into

$$\begin{cases} x'(t) = -a(t)x(t) + \sum_{i=1}^m f_i(t, x(t - \tau_i(t))), & t \neq t_k, \\ x(t_k^+) = (1 + b_k)x(t_k), & k \in \mathbb{N} \end{cases}$$

where:

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where:

- ▶ for  $t \geq 0$  and  $u \geq -y^*(t - \tau_i(t))$ ,  $f_i(t, u) = \beta_i(t)g_i(t, u)$  with

$$g_i(t, u) = \frac{1}{1 + [u + y^*(t - \tau_i(t))]^n} - \frac{1}{1 + y^*(t - \tau_i(t))^n}, \quad (8)$$

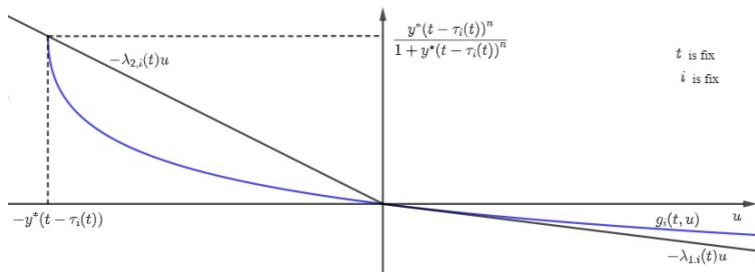
$$S = \left\{ \varphi \in PC : \varphi(\theta) \geq -y^*(\theta) \text{ for } \theta \in [-\bar{\tau}, 0), \varphi(0) > -y^*(0) \right\}$$

Let  $n \in (0, 1]$ .

Considering  $g_i(t, u)$  defined in (8), we have

$$\frac{\partial g_i}{\partial u}(t, u) < 0 \text{ and } \frac{\partial^2 g_i}{\partial u^2}(t, u) > 0, \quad \forall u > -y^*(t - \tau_i(t)), \forall t \geq 0$$

$$\text{with } \frac{\partial g_i}{\partial u}(t, 0) = -\frac{ny^*(t - \tau_i(t))^{n-1}}{[1 + y^*(t - \tau_i(t))^n]^2}.$$



$$\lambda_{1,i}(t) = \frac{ny^*(t - \tau_i(t))^{n-1}}{[1 + y^*(t - \tau_i(t))^n]^2},$$

$$\lambda_{2,i}(t) = \frac{y^*(t - \tau_i(t))^{n-1}}{1 + y^*(t - \tau_i(t))^n}.$$

- **Theorem 3:** Assume **(H1)-(H4)** and  $n \in (0, 1]$ .

The periodic solution  $y^*(t)$  of (5) is GAS, in the set of positive solutions, if there is  $T > 0$  such that

$$\alpha_1^* \alpha_2^* < 1 \quad \text{or} \quad \alpha_1 \alpha_2 < \frac{9}{2},$$

where  $\alpha_j^* = \sup_{t \geq T} \alpha_j^*(t)$ ,  $\alpha_j = \sup_{t \geq T} \alpha_j^*(t) e^{\int_{t-\tau(t)}^t a(u) du}$  ( $j = 1, 2$ ),

and

$$\alpha_1^*(t) = \int_{t-\tau(t)}^t \sum_{i=1}^m \beta_i(s) \frac{ny^*(s - \tau_i(s))^{n-1}}{[1 + y^*(s - \tau_i(s))^n]^2} B_i(s) e^{-\int_s^t a(u) du} ds$$

$$\alpha_2^*(t) = \int_{t-\tau(t)}^t \sum_{i=1}^m \beta_i(s) \frac{y^*(s - \tau_i(s))^{n-1}}{1 + y^*(s - \tau_i(s))^n} B_i(s) e^{-\int_s^t a(u) du} ds$$

$$\text{with } B_i(s) = \prod_{k: t-\tau_i(t) \leq t_k < t} (1 + b_k)^{-1}, \quad i = 1, \dots, m.$$



Let  $n \in (1, \infty)$ .

In this case,  $g_i(t, u)$  defined in (8) verifies

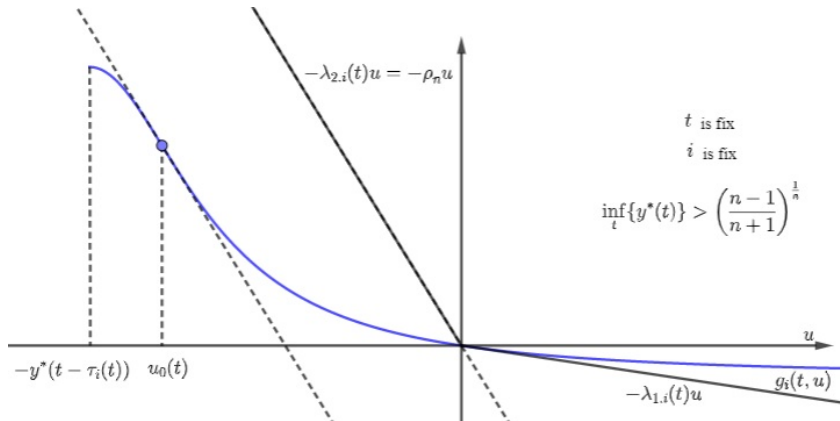
$$\frac{\partial g_i}{\partial u}(t, u) < 0, \quad \forall u > -y^*(t - \tau_i(t)), \quad \forall t \geq 0,$$

and

$$\begin{cases} \frac{\partial^2 g_i}{\partial u^2}(t, u) > 0 \text{ for } u > u_0(t), \\ \frac{\partial^2 g_i}{\partial u^2}(t, u) < 0 \text{ for } u \in (-y^*(t - \tau_i(t)), u_0(t)), \end{cases}$$

where  $u_0(t) := -y^*(t - \tau_i(t)) + \left(\frac{n-1}{n+1}\right)^{\frac{1}{n}}$  is the unique inflection point of  $u \mapsto g_i(t, u)$ .

We have  $\frac{\partial g_i}{\partial u}(t, u_0(t)) = -\rho_n = -\frac{(n+1)^2}{4n} \left(\frac{n-1}{n+1}\right)^{\frac{n-1}{n}}$ .



**Theorem 4:** Assume **(H1)**-**(H4)** and  $n > 1$ .

The periodic solution  $y^*(t)$  of (5) is GAS (in  $PC_0^+$ ) if, for some  $T > 0$ , one of the following conditions holds:

- (i)  $(\alpha_1 \gamma < \frac{9}{4} \text{ or } \alpha_1^* \gamma^* < 1)$  and  $\inf_t \{y^*(t)\} \geq \left(\frac{n-1}{n+1}\right)^{\frac{1}{n}}$ ;
- (ii)  $(\alpha_1 \gamma < \frac{9}{4} \text{ or } \alpha_1^* \gamma^* < 1)$  and  $\sup_t \{y^*(t)\} \leq \left(\frac{n-1}{n+1}\right)^{\frac{1}{n}}$ ;
- (iii)  $\gamma < \frac{3}{2} \text{ or } \gamma^* < 1$ ,

where  $\gamma^* = \sup_{t \geq T} \gamma^*(t)$ ,  $\gamma = \sup_{t \geq T} \gamma^*(t) e^{\int_{t-\tau(t)}^t a(u) du}$ , with

$$\gamma^*(t) = \rho_n \int_{t-\tau(t)}^t \sum_{i=1}^m \beta_i(s) B_i(s) e^{-\int_s^t a(u) du} ds,$$

with  $\rho_n = \frac{(n+1)^2}{4n} \left(\frac{n-1}{n+1}\right)^{\frac{n-1}{n}}$ ,  $B_i(s)$ ,  $\alpha_1$ , and  $\alpha_1^*$  as above.

In case that  $y^*(t)$  is unknown, we have the estimate

$$\mathfrak{m} \leq y^*(t) \leq \mathfrak{M}, \quad t \geq 0,$$

where

$$\mathfrak{M} = \min \left\{ M\beta\bar{B}, M\bar{B}(e^{A(\omega)} - 1)e^{A(\omega)} \left( \max_{t \in [0, \omega]} \frac{\sum_{i=1}^m \beta_i(t)}{a(t)} \right) \right\}$$

$$\mathfrak{m} = \frac{e^{-A(\omega)} M\bar{B}}{1 + \mathfrak{M}^n} \max \left\{ \beta, (e^{A(\omega)} - 1) \left( \min_{t \in [0, \omega]} \frac{\sum_{i=1}^m \beta_i(t)}{a(t)} \right) \right\}$$

with  $\beta = \int_0^\omega \sum_{i=1}^m \beta_i(s) ds$ ,  $A(\omega) = \int_0^\omega a(u) du$ ,

$$M = (\prod_{k=1}^p (1 + b_k)^{-1} - e^{-A(\omega)})^{-1},$$

$$\bar{B} = \max \left\{ 1, \prod_{k=j}^{j+l} (1 + b_k)^{-1} : j = 1, \dots, p, l = 0, \dots, p-1 \right\}, \text{ and}$$

$$\underline{B} = \min \left\{ 1, \prod_{k=j}^{j+l} (1 + b_k)^{-1} : j = 1, \dots, p, l = 0, \dots, p-1 \right\}.$$

- One delay multiple of the period ( $m = 1$ ,  $\tau(t) = q\omega$ ,  $q \in \mathbb{N}$ )

$$\begin{cases} y'(t) = -a(t)y(t) + \frac{\beta(t)}{1 + y(t - q\omega)^n}, & 0 \leq t \neq t_k, \\ y(t_k^+) = (1 + b_k)y(t_k), & k \in \mathbb{N} \end{cases} \quad (9)$$

- ▶ One delay multiple of the period ( $m = 1$ ,  $\tau(t) = q\omega$ ,  $q \in \mathbb{N}$ )

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- ▶ **Theorem 5:** Assume **(H2)**-**(H4)** and  $a, \beta : [0, \infty) \rightarrow (0, \infty)$   $\omega$ -periodic continuous functions.

Then (9) has a positive periodic solution which is GAS if:  
case  $n > 1$

$$\rho_n B_0^q \sup_{t \geq T} \int_{t-q\omega}^t \beta(s) e^{-\int_s^t a(u) du} ds < \max \left\{ 1, \frac{3}{2} e^{-q \int_0^\omega a(u) du} \right\},$$

or, case  $0 < n \leq 1$ ,

$$\sqrt{n} B_0^q \sup_{t \geq T} \int_{t-q\omega}^t \beta(s) e^{-\int_s^t a(u) du} ds < m \max \left\{ 1, \frac{3}{2} e^{-q \int_0^\omega a(u) du} \right\}$$

where  $B_0 = \prod_{k=1}^p (b_k + 1)^{-1}$ .

- **Remark:** Saker and Alzabut [6] proved the existence of a positive periodic solution of (9) and its GAS assuming (H1), (H3),  $n \in \mathbb{N}$ , the function

$$t \mapsto \prod_{k: t_k \in [0, t)} (1 + b_k) \text{ is } \omega\text{-periodic,} \quad (10)$$

and the “3/2-type condition”

$$\rho_n q \int_0^\omega \beta(s) ds < \frac{3}{2} e^{-q \int_0^\omega a(u) du}. \quad (11)$$

[6] S.H. Saker and J.O. Alzabut, *On the impulsive delay hematopoiesis model with periodic coefficients*, Rocky

Mountain J. Math. 39 (2009) 1657-1688.

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Thus Theorem 5 improves the stability criterion in [6].

[6] S.H. Saker and JO. Alzabut, *On the impulsive delay hematopoiesis model with periodic coefficients*, Rocky

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- No impulsive case ( $b_k = 0, \forall k \in \mathbb{N}$ )

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where  $\mathcal{A} = \sup_{t \in [0, \omega]} \int_{t-\tau(t)}^t a(u) du$ , then there is a positive  $\omega$ -periodic solution of (12) which is GAS.

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- ▶ Liu et al [7] proved the existence of a positive periodic solution of (12) and its GAS assuming (H1),  $n > 1$ , and

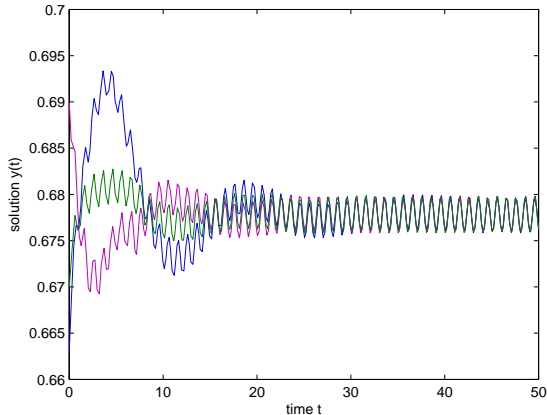
$$(n-1)^{\frac{n-1}{n}} \frac{e^{A(\omega)}}{e^{A(\omega)} - 1} \int_0^\omega \sum_{i=1}^m \beta_i(s) ds \leq 1.$$

# Numerical example

Consider the 1-periodic model

$$y'(t) = - \left( 1 + \frac{1}{2} \cos(2\pi t) \right) y(t) + \frac{\eta_1 \left( 1 + \frac{1}{2} \cos(2\pi t) \right)}{1 + y(t - 6 - \cos(2\pi t))^n} \\ + \frac{\eta_2 \left( 1 + \frac{1}{2} \sin(2\pi t) \right)}{1 + y(t - 7 - \cos(2\pi t))^n} + \frac{\eta_3 \left( 1 + \frac{1}{2} \cos(2\pi t) \right)}{1 + y(t - 15 - \cos(2\pi t))^n},$$

where  $\eta_1, \eta_2, \eta_3$  are positive real numbers.



**Figure:** Numerical simulation of three solutions where  $\eta_1 = 1.1$ ,  $\eta_2 = 0.03$ ,  $\eta_3 = 0.001$  and  $n = 1.03$ , with initial condition  $\varphi(\theta) = 0.67$ ,  $\varphi(\theta) = 0.65(1 + 0.02 \cos(\theta))$ , and  $\varphi(\theta) = 0.69(1 + 0.02 \sin(\theta))$ , for  $\theta \in [-16, 0]$ , respectively.

# Thank you

The presented results are published in

[8] T. Faria and J.J. Oliveira, *Global asymptotic stability for a periodic delay hematopoiesis model with impulses*, Applied Mathematical Modelling 79 (2020) 843-864.