

POSITIVE ALMOST PERIODIC SOLUTIONS FOR THE HEMATOPOIESIS MODEL VIA THE HILBERT PROJECTIVE METRIC

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Abstract

The aim of this work is to prove the existence of a positive almost periodic solution to a multifinite time delayed nonlinear differential equation that describes the so-called hematopoiesis model. The approach uses the Hilbert projective metric in a cone. With some additional assumptions, we construct a fixed point theorem to prove the desired existence and uniqueness of the solution.

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1. Introduction

To describe some physiological control systems in the classic study of population dynamics, Mackey and Glass [8] proposed the following autonomous nonlinear delay differential equation

$$\rho'(t) = -\gamma\rho(t) + \frac{\beta\rho^m(t-\tau)}{1 + \rho^n(t-\tau)}, \quad (1.1)$$

where $\gamma, \beta, n \in (0, +\infty)$, $\tau, m \in [0, +\infty)$, $\rho(t)$ denotes the density of mature blood cells circulating in the bloodstream and γ is the rate of loss of blood cells from the circulation. Here, $f(\rho(t-\tau)) = \beta\rho^m(t-\tau)/(1 + \rho^n(t-\tau))$ is the flux of blood cells into the bloodstream from the stem cell compartment and depends on the delayed density, $\rho(t-\tau)$, of mature cells in circulation, where τ is the time delay between the production of the immature cells in the bone marrow and the release of the mature cells into the bloodstream.

In [9, 13], the model (1.1) is extended to the following nonautonomous nonlinear delay differential equation with time-varying coefficients and delays, which takes more account of real phenomena, such as the important role played by variations in the

environment: namely,

$$x'(t) = -a(t)x(t) + \sum_{i=1}^k \frac{b_i(t)x^m(t - \tau_i(t))}{1 + x^n(t - \tau_i(t))}, \quad (1.2)$$

where $0 \leq m < n$, the functions $a, b_i, \tau_i : \mathbb{R} \rightarrow (0, +\infty)$ are continuous for $i = 1, 2, \dots, k$, $x(t)$ is the density of mature blood cells circulating in the bloodstream and $a(t)$ is the rate of loss of blood cells from the circulation at time t . Now, $f(x(t - \tau_i(t))) = b_i(t)x^m(t - \tau_i(t))/(1 + x^n(t - \tau_i(t)))$ is the flux of blood cells into the bloodstream from the i th stem cell compartment for $i = 1, 2, \dots, k$ and $\tau_i(t)$ is the corresponding time delay between the production of immature cells in the bone marrow in the i th compartment and the release of the mature cells into the bloodstream. The model (1.1) and its extension (1.2) are referred to as a model of hematopoiesis (cell production).

The existence of periodic and almost periodic solutions to the hematopoiesis model has been investigated extensively (see, for example, [4, 7, 12, 14, 15] and references therein). Some interesting results concerning the existence of almost periodic solutions to the hematopoiesis model (1.2) were obtained by Zhang *et al.* using the contraction mapping principle in the case $m = 0$ [14] and by using some additional conditions in the cases $m = 0$ or $m = 1$ [12, 14, 15]. For $0 \leq m \leq 1$ and assuming an *a priori* estimate that controls the balance between the loss rate and the flux, Liu [7] proved the existence of a positive almost periodic solution to (1.2). Recently, Diagana *et al.* [4] established the existence of a positive almost periodic solution to (1.2) via a fixed-point theorem in a cone.

Our purpose in this work is also to give criteria for the existence and uniqueness of a positive almost periodic solution to (1.2). By contrast with Diagana *et al.* [4], we use here the Hilbert projective metric in a cone, which allows the interior to accommodate a complete metric space structure. We follow the development presented in [5].

This paper is organised as follows. In the next section, we present some preliminaries that will be used to prove the main result. Section 3 deals with the main result on the existence and uniqueness of a positive almost periodic solution to (1.2) and the last section gives a concrete illustration of our result.

2. Preliminaries

We recall some definitions, notation and lemmas, which will be used later.

DEFINITION 2.1 [2, 10]. Let X be a Banach space. A continuous function $f : \mathbb{R} \rightarrow X$ is called almost periodic if, for each $\epsilon > 0$, there exists $l(\epsilon) > 0$ such that every interval of length $l(\epsilon)$ contains a number τ with the property that $\|f(t + \tau) - f(t)\| \leq \epsilon$ for each $t \in \mathbb{R}$. The number τ is called an ϵ -translation number of $f(t)$. Denote the set of such functions by $AP(\mathbb{R})$.

LEMMA 2.2. *The intersection of two relatively dense subsets of \mathbb{R} is relatively dense.*

PROOF. Let \mathcal{T} and \mathcal{T}' be two relatively dense subsets of \mathbb{R} and put $\tilde{\mathcal{T}} = \mathcal{T} \cap \mathcal{T}'$. For all $\epsilon > 0$, there exist $L(\mathcal{T}, \epsilon)$ and $L(\mathcal{T}', \epsilon)$ such that $\mathcal{T} \cap [t, t + L(\mathcal{T}, \epsilon)] \neq \emptyset$ and $\mathcal{T}' \cap [t, t + L(\mathcal{T}', \epsilon)] \neq \emptyset$ for all $t \in \mathbb{R}$. If $L(\tilde{\mathcal{T}}, \epsilon) = \max(L(\mathcal{T}, \epsilon), L(\mathcal{T}', \epsilon))$, then $\tilde{\mathcal{T}} \cap [t, t + L(\tilde{\mathcal{T}}, \epsilon)] \neq \emptyset$, which achieves the proof. \square

2.1. Exponential dichotomy.

DEFINITION 2.3 [6, 10]. Let $x \in \mathbb{R}^n$ and let $A(t)$ be an $n \times n$ continuous matrix defined on \mathbb{R} . The linear system

$$x'(t) = A(t)x(t) \tag{2.1}$$

admits an exponential dichotomy on \mathbb{R} if there exist positive constants λ, k and a projection P such that the fundamental solution matrix $X(t)$ of (2.1) satisfies

$$\|X(t)PX^{-1}(s)\| \leq \lambda e^{-k(t-s)} \quad \text{for } t \geq s, \quad \|X(t)(I - P)X^{-1}(s)\| \leq \lambda e^{-k(s-t)} \quad \text{for } t \leq s.$$

LEMMA 2.4 [6, 10]. Let $c_i(t)$ be an almost periodic function on \mathbb{R} and let

$$M[c_i] = \lim_{T \rightarrow \infty} \frac{1}{T} \int_t^{t+T} c_i(s) ds > 0 \quad \text{for } i = 1, 2, \dots, n.$$

Then the linear system $x'(t) = C(t)x(t)$ admits an exponential dichotomy on \mathbb{R} , where $C(t) = \text{diag}(-c_1(t), -c_2(t), \dots, -c_n(t))$.

LEMMA 2.5 [1]. Let f be an almost periodic function. If the linear system (2.1) admits an exponential dichotomy, then the almost periodic system

$$x'(t) = A(t)x(t) + f(t)$$

has a unique almost periodic solution $x(t)$ and

$$x(t) = \int_{-\infty}^t X(t)PX^{-1}(s)f(s) ds - \int_t^{+\infty} X(t)(I - P)X^{-1}(s)f(s) ds,$$

where $X(t)$ is the fundamental solution matrix of (2.1).

2.2. The Hilbert projective metric and fixed point theorem in a cone. Let X be a real Banach space. A closed convex set P in X is called a convex cone if the following conditions are satisfied:

- (i) if $x \in P$, then $\lambda x \in P$ for all $\lambda \geq 0$; and
- (ii) if $x \in P$ and $-x \in P$, then $x = 0$.

A partial ordering \leq in X is induced by P : for all $x, y \in X, x \leq y$ if and only if $y - x \in P$. Given $u, v \in P$, define the interval $[u, v] := \{x \in X : u \leq x \leq v\}$.

A cone P is called normal if there exists a constant $k > 0$ such that, for all $x, y \in P, 0 \leq x \leq y$ implies that $\|x\| \leq k\|y\|$, where $\|\cdot\|$ is the norm on X .

If P is now a general cone in a Banach space X and x and y are elements of $P^* = P - 0_X$, we say that x and y are comparable if there exist real numbers $\alpha > 0$ and $\beta > 0$ such that $\alpha x \leq y \leq \beta x$. This defines an equivalence relation on P^* and divides P^*

into disjoint subsets which we call components of P . If x and y are comparable, then we define the numbers $m(y/x)$ and $M(y/x)$ by

$$m(y/x) := \sup\{\alpha > 0 : \alpha x \leq y\} \quad \text{and} \quad M(y/x) := \inf\{\beta > 0 : y \leq \beta x\}.$$

Thompson [11] introduced a metric d defined as follows: if x and y in P^* are comparable, then

$$d(x, y) := \max(\log M(y/x), \log M(x/y)) = \max(\log M(y/x), -\log m(y/x)). \quad (2.2)$$

If C is a component of P , it is easy to see that d gives a metric on C . Moreover, Thompson proved the following results.

THEOREM 2.6 [11]. *Let P be a normal cone in a Banach space X and let C be a component of P . Then C is a complete metric space with respect to the metric d .*

PROPOSITION 2.7 [11]. *Let P be a normal cone in a Banach space X with nonempty interior P° . Then P° is a component of P .*

Thus, the nonempty interior P° of a normal cone P , is a complete metric space with respect to the metric d . We have the following theorem.

THEOREM 2.8 [3]. *Let E be a complete space with respect to the metric d . Suppose there is a mapping f from E into E satisfying*

$$d(f(x), f(y)) \leq \phi(d(x, y)) \quad \text{for all } x \text{ and } y \text{ in } E,$$

where ϕ is a positive nondecreasing function continuous on $[0, +\infty[$, such that $\phi(r) < r$ for all $r > 0$ and $\phi(0) = 0$. Then f has exactly one fixed point in E .

For a bounded continuous function $h(t)$, we introduce the notation

$$h^+ = \sup_{t \in \mathbb{R}} h(t), \quad h^- = \inf_{t \in \mathbb{R}} h(t). \quad (2.3)$$

3. Positive almost periodic solutions

This section contains the proof of our main result on the existence and uniqueness of a positive almost periodic solution of (1.2). The proof is based on a fixed point theorem in a cone endowed with the Hilbert projective metric.

3.1. Assumptions and main result. For $t \in \mathbb{R}$ and $x \in \mathbb{R}^+$, put

$$F(t, x) = \sum_{i=1}^k b_i(t) f[x(t - \tau_i(t))] = \sum_{i=1}^k \frac{b_i(t) x^m(t - \tau_i(t))}{1 + x^n(t - \tau_i(t))}.$$

We make two sets of assumptions. The first gives some preliminary estimates and the second deals with the behaviour of flux term. Here, a^+ , a^- , b^+ and b^- are as in (2.3).

(H1) $a^- > 0, b_i^- > 0, \tau_i \geq 0$ for $i = 1, 2, \dots, m$.

(H2) $(\sum_{i=1}^k b_i^+) / a^- \leq (n/(n - m))(m/(n - m))^{(1-m)/n}$.

(H3) $(\sum_{i=1}^k b_i^-)/a^+ > 1$, that is, the flux is always greater than the loss of blood cells in circulation.

(H4) τ_i and b_i for $i = 1, 2, \dots, k$ are almost periodic.

Define

$$f(x) = \begin{cases} \frac{x^m}{1 + x^n} & \text{for } 0 \leq x \leq \left(\frac{m}{n-m}\right)^{1/n}, \\ \frac{n-m}{n} \left(\frac{m}{n-m}\right)^{m/n} & \text{for } x > \left(\frac{m}{n-m}\right)^{1/n}. \end{cases} \tag{3.1}$$

The following theorem is our main result.

THEOREM 3.1. *Suppose that (H1)–(H4) hold. Then (1.2) has exactly one positive almost periodic solution.*

3.2. Proof of the main result.

LEMMA 3.2. *Suppose that φ and σ are in $AP(\mathbb{R})$. Then the function $t \mapsto \varphi(t - \sigma(t))$ is also in $AP(\mathbb{R})$.*

PROOF. Let $\epsilon > 0$ and let τ be a common almost period for φ and σ : that is $\tau \in \tilde{\mathcal{T}}(\varphi, \sigma, \epsilon) = \tilde{\mathcal{T}}(\varphi, \epsilon) \cap \tilde{\mathcal{T}}(\sigma, \epsilon)$, where $\tilde{\mathcal{T}}(\varphi, \epsilon)$ and $\tilde{\mathcal{T}}(\sigma, \epsilon)$ are the sets of ϵ -almost periods associated, respectively, to φ and σ . Note that $\tilde{\mathcal{T}}(\varphi, \sigma, \epsilon) = \tilde{\mathcal{T}}(\varphi, \epsilon) \cap \tilde{\mathcal{T}}(\sigma, \epsilon)$ is relatively dense in \mathbb{R} , by Lemma 2.2. Consider

$$\begin{aligned} |\varphi(t - \sigma(t)) - \varphi(t + \tau - \sigma(t + \tau))| &\leq |\varphi(t - \sigma(t)) - \varphi(t - \sigma(t) + \tau)| \\ &\quad + |\varphi(t + \tau - \sigma(t)) - \varphi(t + \tau - \sigma(t + \tau))|. \end{aligned}$$

Let $\tilde{\epsilon} > 0$. By the uniform continuity of φ , there exists $\epsilon > 0$ such that, for all $t \in \mathbb{R}$, $|\sigma(t) - \sigma(t + \tau)| \leq \epsilon$ implies that $|\varphi(t + \tau - \sigma(t)) - \varphi(t + \tau - \sigma(t + \tau))| \leq \tilde{\epsilon}$. By the almost periodicity of φ , also $|\varphi(t - \sigma(t)) - \varphi(t - \sigma(t) + \tau)| \leq \epsilon$. Thus we deduce the almost periodicity of $t \mapsto \varphi(t - \sigma(t))$. \square

LEMMA 3.3. *Suppose (H1), (H2) and (H4) hold. Then (1.2) has a nonnegative almost periodic solution x which is given for $t \in \mathbb{R}$ by*

$$x(t) = \int_{-\infty}^t g(s) \sum_{i=1}^k b_i(s) f(x(s - \tau_i(s))) ds \quad \text{where } g(s) = \exp\left(-\int_{-s}^t a(r) dr\right). \tag{3.2}$$

In fact, every nonnegative almost periodic solution φ of (1.2) is also a nonnegative almost periodic solution of (3.2) and vice versa.

PROOF. If φ is a positive almost periodic solution of (1.2), then, by hypothesis (H4) and Lemma 3.2, $\varphi(\cdot - \tau_i(\cdot))$ is almost periodic for $i = 1, 2, \dots, k$. Therefore, the function $\sum_{i=1}^k b_i(\cdot) \varphi^m(\cdot - \tau_i(\cdot)) / (1 + \varphi^n(\cdot - \tau_i(\cdot))) \in AP(\mathbb{R})$. Since $a^- > 0$, from (H1), Lemmas 2.4 and 2.5 yield

$$\varphi(t) = \int_{-\infty}^t g(s) \sum_{i=1}^k \frac{b_i(s) \varphi^m(s - \tau_i(s))}{1 + \varphi^n(s - \tau_i(s))} ds \quad \text{for } t \in \mathbb{R}.$$

Note that

$$\sup_{x \geq 0} \frac{x^m}{1 + x^n} = \frac{n - m}{n} \left(\frac{m}{n - m} \right)^{m/n}.$$

So, by **(H2)**,

$$\begin{aligned} \varphi(t) &\leq \int_{-\infty}^t e^{-a^-(t-s)} \left(\sum_{i=1}^k b_i^+ \frac{n - m}{n} \left(\frac{m}{n - m} \right)^{m/n} \right) ds \\ &= \frac{\sum_{i=1}^k b_i^+}{a^-} \frac{n - m}{n} \left(\frac{m}{n - m} \right)^{m/n} \leq \left(\frac{m}{n - m} \right)^{1/n}. \end{aligned}$$

By (3.1), $f(s - \tau - i(s)) = \varphi^m(s - \tau_i(s)) / (1 + \varphi^n(s - \tau_i(s)))$ for $s \in \mathbb{R}$ and $i = 1, 2, \dots, k$. Thus,

$$\varphi(t) = \int_{-\infty}^t g(s) \sum_{i=1}^k b_i(s) f(\varphi(s - \tau_i(s))) ds \quad \text{for } t \in \mathbb{R}$$

is an almost periodic solution of (3.2).

Similarly, we can show that every nonnegative almost periodic solution φ of (3.2) is also an almost periodic solution of (1.2). \square

In the subsequent work, $Q = \{x \in AP(\mathbb{R}) : x(t) \geq 0, \text{ for all } t \in \mathbb{R}\}$ denotes the normal solid cone in $AP(\mathbb{R})$ and $Q^\circ = \{x \in AP(\mathbb{R}) : \text{there is } \epsilon > 0 \text{ such that } x(t) > \epsilon, \text{ for all } t \in \mathbb{R}\}$ denotes its interior. Let \mathcal{T} be an operator on Q° defined by

$$\begin{aligned} \mathcal{T}(x)(t) &= \int_{-\infty}^t g(s) \sum_{i=1}^k b_i(s) f[x(s - \tau_i(s))] ds \quad \text{for } t \in \mathbb{R}, \\ \text{where } g(s) &= \exp\left(-\int_{-s}^t a(r) dr\right). \end{aligned} \tag{3.3}$$

PROPOSITION 3.4. \mathcal{T} maps Q° into itself.

PROOF. Let x be in Q° . By Lemma 3.3, $\mathcal{T}(x)$ is an almost periodic function. In addition, there exists $\epsilon_0 > 0$ such that $x(t) \geq \epsilon_0$ for all $t \in \mathbb{R}$. Thus

$$\begin{aligned} \mathcal{T}(x)(t) &\geq \int_{-\infty}^t e^{-a^+(t-s)} \sum_{i=1}^k b_i^- \cdot \min\left\{ \frac{\epsilon_0^m}{1 + \epsilon_0^n}, \frac{n - m}{n} \left(\frac{m}{n - m} \right)^{m/n} \right\} ds \\ &= \frac{\sum_{i=1}^k b_i^-}{a^+} \cdot \min\left\{ \frac{\epsilon_0^m}{1 + \epsilon_0^n}, \frac{n - m}{n} \left(\frac{m}{n - m} \right)^{m/n} \right\} = \frac{\sum_{i=1}^k b_i^-}{a^+} \cdot \frac{\epsilon_0^m}{1 + \epsilon_0^n}. \end{aligned}$$

By **(H3)**, $\mathcal{T}(x)(t) > 0$ for all $t \in \mathbb{R}$, which implies that $\mathcal{T}(x) \in Q^\circ$. \square

Next, we will prove the fixed point theorem for the operator \mathcal{T} .

PROPOSITION 3.5. \mathcal{T} is a nondecreasing operator on Q° .

PROOF. Choose x and y in Q° such that $x(t) \geq y(t)$ for all $t \in \mathbb{R}$ and define $g(s)$ as in (3.3). Then, for all $t \in \mathbb{R}$,

$$\mathcal{T}(x)(t) - \mathcal{T}(y)(t) = \int_{-\infty}^t g(s) \sum_{i=1}^k b_i(s)(f[x(s - \tau_i(s))] - f[y(s - \tau_i(s))]) ds.$$

From (3.1), f is nondecreasing on $[0, (m/(n - m))^{1/n}]$ and remains constant on $[(m/(n - m))^{1/n}, \infty)$. Therefore, $\mathcal{T}(x)(t) - \mathcal{T}(y)(t) \geq 0$ for all $t \in \mathbb{R}$. \square

PROPOSITION 3.6. Define the metric d as in (2.2). There exists a positive nondecreasing function ϕ defined on \mathbb{R}^+ , satisfying

$$\phi(0) = 0, \quad \phi(r) < r \quad \text{for } r > 0 \quad \text{and} \quad d(\mathcal{T}(x), \mathcal{T}(y)) \leq \phi(d(x, y)) \quad \text{for } x, y \in Q^\circ.$$

PROOF. Let x and y be two comparable functions in Q° and let $\alpha = m(y/x), \beta = M(y/x)$. Then $\alpha x \leq y \leq \beta x$ and, from (2.2), $d(x, y) = \max(\log(\beta), -\log(\alpha))$. By Proposition 3.5, the operator \mathcal{T} is nondecreasing and so

$$\mathcal{T}(\alpha x) \leq \mathcal{T}(y) \leq \mathcal{T}(\beta x). \tag{3.4}$$

Thus, we have the following cases.

Case 1. $\beta \in (0, 1)$. Then $\alpha \in (0, 1)$ and $f(\alpha x) \geq \alpha^n f(x)$. Therefore, $\mathcal{T}(\alpha x) \geq \alpha^m \mathcal{T}(x)$ and the left-hand side of (3.4) gives $\varphi(\alpha)\mathcal{T}(x) \leq \mathcal{T}(y)$, where $\varphi(\alpha) = \alpha^m$. For the right-hand side (3.4), consider the nondecreasing function

$$\chi(x) = \frac{x^n}{1 + \beta^n x^n} \quad \text{for } 0 < x \leq \left(\frac{m}{n - m}\right)^{1/n},$$

which attains its maximum $\psi(\beta) = [(n/m)(1 - \beta^n) - 1]^{-1}$ when $x = (m/(n - m))^{1/n}$. Then,

$$\frac{f(\beta x)}{f(x)} = \beta^m \frac{1 + x^n}{1 + \beta^n x^n} \leq \beta^m [1 + \chi(x)] \leq \psi(\beta)$$

and we conclude that $\mathcal{T}(y) \leq \psi(\beta)\mathcal{T}(x)$. Therefore, $\alpha x \leq y \leq \beta x$ implies that

$$\varphi(\alpha)\mathcal{T}(x) \leq \mathcal{T}(y) \leq \psi(\beta)\mathcal{T}(x). \tag{3.5}$$

Note from (3.5) that, for all $\alpha, \beta \in (0, 1)$ with $\alpha \leq \beta$, $\varphi(\alpha) \leq \psi(\beta)$. To compute the metric, note that $M(\mathcal{T}(x), \mathcal{T}(y)) \leq \psi(\beta)$ and $m(\mathcal{T}(x), \mathcal{T}(y)) \geq \varphi(\alpha)$, and hence

$$d(\mathcal{T}(x), \mathcal{T}(y)) \leq \max(\log(\Psi(\beta)), -\log(\varphi(\alpha))).$$

Define the function ϕ by $\phi(0) = 0$ and, for $u > 0$,

$$\phi(u) = \max[-\log(\varphi(e^{-u})), \log(\psi(e^u))]. \tag{3.6}$$

Then ϕ is a nondecreasing function and we obtain

$$\begin{aligned} \phi(-\log(\alpha)) &= \max[-\log(\varphi(e^{-(-\log \alpha)})), \log(\psi(e^{-\log \alpha}))] \\ &= \max[-\log(\varphi(\alpha)), \log(\psi(\alpha^{-1}))] \end{aligned}$$

and

$$\begin{aligned}\phi(\log(\beta)) &= \max[-\log(\varphi(e^{-\log\beta})), \log(\psi(e^{\log\beta}))] \\ &= \max[-\log(\varphi(\beta^{-1})), \log(\psi(\beta))].\end{aligned}$$

Thus,

$$d(\mathcal{T}(x), \mathcal{T}(y)) \leq \phi(d(x, y)).$$

Case 2. $\alpha > 1$. Rewrite $\alpha x \leq y \leq \beta x$ as $\beta^{-1}y \leq x \leq \alpha^{-1}y$ and apply Case 1. This yields

$$\psi^{-1}(\alpha^{-1})\mathcal{T}(x) \leq \mathcal{T}(y) \leq \varphi^{-1}(\beta^{-1})\mathcal{T}(x).$$

Next, $M(\mathcal{T}(x), \mathcal{T}(y)) \leq \varphi^{-1}(\beta^{-1})$ and $m(\mathcal{T}(x), \mathcal{T}(y)) \geq \psi^{-1}(\alpha^{-1})$, so that

$$d(\mathcal{T}(x), \mathcal{T}(y)) \leq \max(\log(\varphi^{-1}(\beta^{-1})), -\log(\psi^{-1}(\alpha^{-1}))).$$

Define the function ϕ as in (3.6). Then

$$\begin{aligned}\phi(-\log(\alpha)) &= \max[-\log(\varphi(e^{-(-\log\alpha)})), \log(\psi(e^{-\log\alpha}))] \\ &= \max[-\log(\varphi(\alpha)), \log(\psi(\alpha^{-1}))] = \max[\log(\varphi^{-1}(\alpha)), -\log(\psi^{-1}(\alpha^{-1}))]\end{aligned}$$

and

$$\begin{aligned}\phi(\log(\beta)) &= \max[-\log(\varphi(e^{-\log\beta})), \log(\psi(e^{\log\beta}))] \\ &= \max[-\log(\varphi(\beta^{-1})), \log(\psi(\beta))] = \max[\log(\varphi^{-1}(\beta^{-1})), -\log(\psi^{-1}(\beta))].\end{aligned}$$

Thus,

$$d(\mathcal{T}(x), \mathcal{T}(y)) \leq \phi(d(x, y)).$$

Case 3. $\alpha \leq 1$ and $\beta \geq 1$. This case is easily deduced from the previous cases giving

$$\varphi(\alpha)\mathcal{T}(x) \leq \mathcal{T}(y) \leq \varphi^{-1}(\beta^{-1})\mathcal{T}(x).$$

Here, $M(\mathcal{T}(x), \mathcal{T}(y)) \leq \varphi^{-1}(\beta^{-1})$ and $m(\mathcal{T}(x), \mathcal{T}(y)) \geq \varphi(\alpha)$. Thus,

$$d(\mathcal{T}(x), \mathcal{T}(y)) \leq \max(\log(\varphi^{-1}(\beta^{-1})), -\log(\varphi(\alpha))).$$

Define the function ϕ by $\phi(u) = -\log(\varphi(e^{-u}))$ for $u > 0$, and $\phi(0) = 0$. Then $-\log(\varphi(\alpha)) = \phi(-\log(\alpha))$ and $\log(\varphi^{-1}(\beta^{-1})) = \phi(\log(\beta))$ and, by the monotonicity of ϕ , we conclude that

$$d(\mathcal{T}(x), \mathcal{T}(y)) \leq \phi(d(x, y)).$$

This completes the proof of the proposition. \square

PROOF OF THEOREM 3.1. By Proposition 3.6, the operator \mathcal{T} satisfies all assumptions of Theorem 2.8 and so it has exactly one fixed point $z \in Q^\circ$. By Lemma 3.3, this gives the unique almost periodic solution of (1.2). \square

4. Application

The following example illustrates our results. Consider the hematopoiesis model

$$x'(t) = -\frac{1}{2}\left(1 + \frac{1}{2}\cos t\right)x(t) + \frac{1}{2}\left(2 + \frac{1}{2}|\cos \sqrt{2}t|\right)\frac{x^{1/4}(t - 2e^{\sin^2 t})}{1 + x^{1/2}(t - 2e^{\sin^2 t})} + \frac{1}{2}\left(2 + \frac{1}{2}|\sin \sqrt{3}t|\right)\frac{x^{1/4}(t - 2e^{\sin^2 t})}{1 + x^{1/2}(t - 2e^{\sin^2 t})}. \quad (4.1)$$

The first term on the right-hand side, $a(t) = \frac{1}{2}(1 + \frac{1}{2}\cos t)$, represents the loss rate from the circulation, the flux rate is $F(t, x) = (b_1(t) + b_2(t))x^m/(1 + x^n)$, where $m = \frac{1}{4}$, $n = \frac{1}{2}$, $b_1(t) = \frac{1}{2}(2 + \frac{1}{2}|\cos \sqrt{2}t|)$ and $b_2(t) = \frac{1}{2}(2 + \frac{1}{2}|\sin \sqrt{3}t|)$, and the delays are $\tau_1(t) = \tau_2(t) = 2e^{\sin^2 t}$. It follows that

$$\lim_{T \rightarrow \infty} \frac{1}{T} \int_t^{t+T} a(s) ds = \frac{1}{2},$$

which implies that the equation

$$x'(t) = -\frac{1}{2}\left(1 + \frac{1}{2}\cos t\right)x(t) \quad t \in \mathbb{R},$$

has an exponential dichotomy. We know that $a^+ = \frac{3}{4}$, $a^- = \frac{1}{4}$ and, for $i = 1, 2$, $b_1^- = b_2^- = 1$ and $\tau_i \geq 0$, so hypothesis **(H1)** holds. Also, $b_1^+ = b_2^+ = \frac{5}{4}$, so hypotheses **(H2)** and **(H3)** hold as well. For $i = 1, 2$, b_i and τ_i are almost periodic, so hypothesis **(H4)** holds. In addition, from (3.1), $F(t, x)$ is a nondecreasing function in x and, since $m = \frac{1}{4} < n = \frac{1}{2}$, the density of cells in the blood is always less than one (that is, $x \leq 1$). Consequently, we can apply Theorem 3.1 to this example.

THEOREM 4.1. *The Hematopoiesis model (4.1) has a unique nonnegative almost periodic solution.*

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