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# Mathematical modeling of malaria transmission global dynamics: taking into account the immature stages of the vectors

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## Abstract

In this paper we present a mathematical model of malaria transmission. The model is an autonomous system, constructed by considering two models: a model of vector population and a model of virus transmission. The threshold dynamics of each model is determined and a relation between them established. Furthermore, the Lyapunov principle is applied to study the stability of equilibrium points. The common basic reproduction number has been determined using the next generation matrix and its implication for malaria management analyzed. Hence, we show that if the threshold dynamics quantities are less than unity, the mosquitoes population disappears leading to malaria disappearance; but if they are greater than unity, mosquitoes population persists and malaria also.

Finally, numerical simulations are carried out to support our mathematical results.

**Keywords:** Mosquitoes; Malaria transmission; Thresholds dynamics; Stability; Lyapunov principle

## **1** Introduction

The burden of infectious diseases goes beyond the individual but extends to collectives including families, communities, countries and the whole world. The impact is both social and economic as it keeps children away from school and adults away from work. Most malaria-related mortality and a large fraction of malaria cases occur in sub-Saharan Africa, where transmission is very intense. Moreover, in endemic regions, children under five, pregnant women, and non-immune adults are most at risk of mortality due to malaria. For instance, the World Health Organization estimated that there were 214 million malaria cases in 2015, resulting in about 438 thousand deaths. Costs for treatment are often very expensive for patients driving already poor families into ruin. The countrywide economic loss due to disease is immense, cementing poverty and underdevelopment particularly in low income countries [29, 30].

Although mathematical models are an abstract simplification of the reality; they can still capture the main features of the system and are more amenable to experimentation or analysis. As such, mathematical models can therefore be used to describe or predict the outcomes of epidemics or pandemics providing information that is crucial in informing public health intervention policies. This allows policy makers to optimize the use of



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their limited resources. Concerning the mathematical modeling of malaria, significant results have been established in the recent years since the first model introduced by Ross [28]. Ross defended that keeping the mosquitoes population under a certain threshold can lead to malaria eradication. Some years later, Macdonald [22] improved the model of Ross showing that reducing the number of mosquitoes effectively has some effects on epidemiology of malaria in areas of intense transmission. Before the role of anopheles in the spread of malaria was known, efforts to control the disease were sporadic, infrequent and insignificant. Furthermore, Aron and May [2], added various features of malaria to the model of Macdonald, such that an incubation period in the mosquito, super-infection and a period of immunity in human beings. Besides, the inclusion of acquired immunity proposed by Dietz et al. [12] was a major point of malaria modeling.

Other reviews on mathematical modeling in malaria include work by Ngwa et al. [24] and Chitnis et al. [7, 9]. Indeed, in the model proposed by Ngwa and Shu, human hosts follow an SEIRS-like pattern and vector hosts follow the SEI pattern due to their short life cycle. In [35], a similar model is described by Yang, but with only one class for humans. Humans move from the susceptible, to the exposed class at some probability when they come into contact with an infectious mosquito, and then to the infectious class, as in conventional SEIRS models. However, infectious people can then recover with, or without, a gain in immunity; and either return to the susceptible class, or move to the recovered class. Moreover, Chitnis et al. extended the model of Ngwa and Shu by assuming that, although individuals in the recovered class are immune, in the sense that they do not suffer from serious illness and do not contract clinical malaria, they still have low levels of *Plasmod-ium* in their blood stream and can infect the susceptible mosquitoes. This is one of the main features which makes a distinction between malaria and many other vector-borne diseases.

In addition, based on the susceptibility, the exposedness and the infectivity of human hosts, Ducrot et al. [13] have developed two species malaria model in which we find two host types in the human population: non-immune and semi-immune. In fact, the non-immune is supposed to be more vulnerable to malaria than the semi-immune because it has never acquired immunity against the disease. Meanwhile, the semi-immune has at least once acquired immunity in his life.

In the study of all these models, we remark that the mosquito life cycle is ignored. Generally, the authors consider a constant recruitment rate in the vector population. However, recent work has shown that some of the factors, as the age structure of mosquitoes population and climate effects, are very important for a better understanding of malaria transmission global dynamics, [5, 8, 13, 14, 23, 25, 31]. Indeed, mosquitoes undergo complete metamorphosis going through four distinct stages of development during a lifetime: egg, larva, pupa, and adult. While it is appropriate to assume that only adult mosquito are involved in the malaria transmission, the dynamics of the juvenile stages (larvae and pupae) has significant effects on the dynamics of the mosquitoes population, and then the malaria transmission global dynamics. Motivated by this work, and using the malaria model in [13] as our baseline model, we include the four distinct metamorphic stages of mosquito to formulate a mosquito-stage-structured autonomous model of malaria spread in a more general setting.

The paper is organized as follows. In Sect. 2, we present a vector age-structured model. For this model we established a threshold parameter *r*. Using this threshold and the Lya-

punov theory, we established the local stability and the global stability of the equilibria. Section 3 concerns the malaria transmission model. Its mathematical analysis is done focusing on the boundedness, the positivity, local stability, and global stability of the equilibria. We also found two threshold parameters, respectively, denoted by r and  $r_0$ , that determine the global dynamical behavior of malaria in an area. Section 4 is devoted to numerical simulations. A conclusion finishes the paper.

### 2 Vectors population growth dynamics

## 2.1 Mathematical formulation of the model

In this section, we formulate a model for the mosquitoes population growth basing on their life cycle. There are four main stages in the vector life cycle. The first three stages namely egg, larva and pupa are both aquatics, while the adult stage is aerial. Moreover, the eggs, the larvae and the pupae respond differently to the control measures. For instance, chemical interventions on the breeding sites has impact on the larvae and pupae population, but not on the eggs. For all these reasons, to provide some acceptable strategies to stop mosquitoes population proliferation, it would be fair to dissociate the aquatic stage in the modeling of the mosquitoes growth dynamics. Here, we propose a mathematical model of female anopheles population global behavior as in [1, 23] taking into account the four stages of mosquito. Thus, following the four different stages of the mosquito growth dynamics, we, respectively, denote

- *E*(*t*): the number of eggs at the moment *t*;
- *L*(*t*): the number of larvae at the moment *t*;
- *P*(*t*): the number of pupae at the moment *t*;
- *A*(*t*): the number of females at the moment *t*.

Let us consider the following positive transfer parameters:

- *b*: the intrinsic egg-laying rate;
- *s<sub>E</sub>*, *s<sub>L</sub>*, *s<sub>P</sub>*: respectively, the rates of transfer from eggs to larvae, from larvae to pupae, and from pupae stage to females.
- $d_E$ ,  $d_L$ ,  $d_P$ ,  $f_m$ : respectively, the natural death rate of eggs, larvae, pupae, and females.
- (H1): We assume that the number of eggs is proportional to the number of females.

Using the hypothesis (H1) and analyzing the above diagram (Fig. 1), we obtain the following system:

$$\begin{cases} E'(t) = bA(t) - (s_E + d_E)E(t), \\ L'(t) = s_E E(t) - (s_L + d_L)L(t), \\ P'(t) = s_L L(t) - (s_P + d_P)P(t), \\ A'(t) = s_P P(t) - f_m A(t). \end{cases}$$
(1)

In this model, we only take into account the phenomena of growth and death of the different stages.

However, when we consider the difficulties due to the availability of spaces, foods and the oviposition habitat selection, the following assumptions can be made:

(H2): The growth of eggs depends on the availability of the nutrients. It also depends on the availability of the space because, the oviposition habitat selection is made taking into account the possibility of development of larvae and pupae. Indeed, before



laying eggs the adult mosquitoes make sure that the immature stages can develop relatively unmolested. This hypothesis leads to the following logistic growth model:

$$\begin{cases} E'(t) = bA(t)(1 - \frac{E(t)}{K_E}) - (s_E + d_E)E(t), \\ L'(t) = s_E E(t)(1 - \frac{L(t)}{K_L}) - (s_L + d_L)L(t), \\ P'(t) = s_L L(t)(1 - \frac{P(t)}{K_P}) - (s_P + d_P)P(t), \\ A'(t) = s_P P(t) - f_m A(t). \end{cases}$$
(2)

## 2.2 Mathematical analysis of the vector model

## 2.2.1 Existence and uniqueness of solutions

**Theorem 2.1** For any initial condition  $(t_0, X_0) \in \mathbb{R}_+ \times \mathbb{R}^4_+$ , the system (2) admits a unique positive maximal solution.

*Proof* The model (2) is described by a system of first order autonomous linear differential equations. It can be rewritten as follows:

$$X'(t) = \mathcal{F}_1(X(t)),$$

where

$$X(t) = \left(E(t), L(t), P(t), A(t)\right)^{T}$$

and  $\mathcal{F}_1$  is  $\mathcal{C}^\infty$  of  $\mathbb{R}^4$  into  $\mathbb{R}^4$  and defined by

$$\mathcal{F}_{1}(X) = \begin{pmatrix} f_{1}(x_{1}, x_{2}, x_{3}, x_{4}) \\ f_{2}(x_{1}, x_{2}, x_{3}, x_{4}) \\ f_{3}(x_{1}, x_{2}, x_{3}, x_{4}) \\ f_{4}(x_{1}, x_{2}, x_{3}, x_{4}) \end{pmatrix} = \begin{pmatrix} bx_{4}(1 - \frac{x_{1}}{K_{E}}) - (s_{E} + d_{E})x_{1} \\ s_{E}x_{1}(1 - \frac{x_{2}}{K_{L}}) - (s_{L} + d_{L})x_{2} \\ s_{L}x_{2}(1 - \frac{x_{3}}{K_{P}}) - (s_{P} + d_{P})x_{3} \\ s_{P}x_{3} - f_{m}x_{4} \end{pmatrix}$$

with  $X = (x_1, x_2, x_3, x_4) \in \mathbb{R}^4$ . Since  $\mathcal{F}_1$  is  $\mathcal{C}^\infty$  and then  $\mathcal{C}^1$ , it is locally lipschitzian on  $\mathbb{R}^4$ . Then we deduce the existence and the uniqueness of the maximal solution of the Cauchy problem associated to the system (2) with the initial condition  $(t_0, X_0) \in \mathbb{R}_+ \times \mathbb{R}^4$ . In addition, the solution is  $\mathcal{C}^\infty$  because  $\mathcal{F}_1$  is  $\mathcal{C}^\infty$ .

Now, we establish the non-negativity of the solutions. For this purpose, we proceed by absurd. Let us assume that there exists  $\bar{t}_1 > t_0$  such that  $\forall t > \bar{t}_1, X(t) \notin \mathbb{R}^4_+$ . Consider

$$t_1 = \inf\{t | X(t) \notin \mathbb{R}^4_+\},\tag{3}$$

this means that  $\forall t \in \mathbb{R}_+, t_0 \leq t < t_1, X(t) \in \mathbb{R}_+^4$ .

Consequently, there exists  $\epsilon > 0$  such that

$$\forall t_1 \le t < t_1 + \epsilon, \quad X(t) \notin \mathbb{R}^4_+. \tag{4}$$

Since  $X^* = (0, 0, 0, 0)$  is a steady equilibrium, the uniqueness of the solutions implies  $X(t_1) \neq (0, 0, 0, 0)$ .

- For  $t = t_1$ , 12 cases are possible.
- (i) Let us consider the case  $X(t_1) = (0, L(t_1), P(t_1), A(t_1))$ , where  $L(t_1), P(t_1), A(t_1)$  are positive. Since  $A(t_1) > 0$  and  $E(t_1) = 0$ , from the first equation of the system (2) we have

$$E'(t_1) = bA(t_1) > 0.$$

A first order limited development of E(t) in the neighborhood of  $t_1$  is given by

$$E(t) = E'(t_1)(t - t_1) + o(t - t_1), \quad t \to t_1.$$

Thus, there exists  $\overline{\epsilon} > 0$  such that, for all  $t \in [t_1, t_1 + \overline{\epsilon}]$ , we have E(t) > 0. Besides, by continuity of solutions, there exists  $\overline{\overline{\epsilon}} > 0$  such that, for all  $t \in [t_1, t_1 + \overline{\overline{\epsilon}}]$ , L(t) > 0, P(t) > 0, A(t) > 0. Thus,

$$t \in [t_1, t_1 + \inf\{\bar{\epsilon}, \bar{\bar{\epsilon}}\}], \text{ and } X(t) \in \mathbb{R}^4_+.$$

This result is a contradiction with the definition of  $t_1$  given in (3). (ii) Let us consider  $X(t_1) = (0, 0, 0, A(t_1))$  with  $A(t_1) > 0$ .

We previously show that there exists  $\overline{\epsilon} > 0$  such that, for all  $t \in [t_1, t_1 + \overline{\epsilon}]$ , L(t) > 0. Besides,  $P(t_1) = 0$ ,  $P'(t_1) = 0$ ,  $P''(t_1) = 0$  and  $P'''(t_1) = s_L L''(t_1)$ . Since  $L''(t_1) = s_E E'(t_1) = s_E b A(t_1) > 0$  for  $A(t_1) > 0$ ,

$$P^{\prime\prime\prime}(t_1) = s_E s_L b A(t_1) > 0.$$

Therefore, a three order limited development of P(t) about  $t_1$  is written as follows:

$$P(t) = P^{\prime\prime\prime}(t_1) \frac{(t-t_1)^3}{6} + o\big((t-t_1)^3\big), \quad t \to t_1.$$

We deduce that there exists  $\overline{\overline{\epsilon}} > 0$  such that, for all  $t \in [t_1, t_1 + \overline{\overline{\epsilon}}]$ , P(t) > 0 and therefore, since  $A(t_1) > 0$ , there exists  $\epsilon_1 > 0$  such that, for all  $t \in [t_1, t_1 + \epsilon_1]$ ,  $X(t) \in \mathbb{R}^4_+$ . This is a contradiction with the condition (4).

A similar proof can easily be given for the ten other cases, which are  $(E(t_1), L(t_1), P(t_1), 0)$ ,  $(E(t_1), L(t_1), 0, 0)$ ,  $(E(t_1), 0, 0, 0)$ ,  $(E(t_1), 0, P(t_1), 0)$ ,  $(E(t_1), 0, 0, A(t_1))$ ,  $(0, L(t_1), P(t_1), 0)$ ,  $(0, L(t_1), 0, A(t_1))$ ,  $(0, 0, P(t_1), 0)$ ,  $(0, L(t_1), 0, 0)$ ,  $(0, 0, P(t_1), A(t_1))$  where  $E(t_1), L(t_1), P(t_1)$  and  $A(t_1)$  are, respectively, positive.

We investigate the asymptotic behavior of orbits starting in the non-negative cone,

$$\mathbb{R}^{4}_{+} = \left\{ (x, y, z, w) \in \mathbb{R}^{4} | x \ge 0, y \ge 0, z \ge 0, w \ge 0 \right\}.$$

Let us also consider the positive cone denoted by

$$\mathbb{R}^{*4}_{+} = \left\{ (x, y, z, w) \in \mathbb{R}^{4} | x > 0, y > 0, z > 0, w > 0 \right\}.$$

#### 2.2.2 Equilibrium states

Let us consider the following threshold parameter, called the mosquito reproduction number:

$$r = \left(\frac{b}{s_E + d_E}\right) \left(\frac{s_E}{s_L + d_L}\right) \left(\frac{s_L}{s_P + d_P}\right) \left(\frac{s_P}{f_m}\right)$$

**Proposition 2.1** The system (2) always has a mosquito-free equilibrium

$$X_0^* = (0, 0, 0, 0).$$

Moreover,

- *if*  $r \leq 1$ , *then system* (2) *has no other equilibrium,*
- *if* r > 1, *there is a unique non-trivial equilibrium*,

$$X^* = \left(1 - \frac{1}{r}\right) \left(\frac{K_E}{\chi_E}, \frac{K_L}{\chi_L}, \frac{K_P}{\chi_P}, \frac{s_P}{f_m} \frac{K_P}{\chi_P}\right) = \left(E^*, L^*, P^*, A^*\right),$$

where

$$\begin{split} \chi_E &= \left(1 - \frac{1}{r}\right) + \frac{f_m(s_E + d_E)\chi_P}{bs_P K_P},\\ \chi_L &= \left(1 - \frac{1}{r}\right) \left(1 + \frac{(s_L + d_L)K_L}{s_E K_E}\right) + \frac{(s_E + d_E)(s_L + d_L)K_L\chi_P}{bs_E s_P K_P}, \end{split}$$

and

$$\chi_P = 1 + \frac{s_E K_E K_P (s_P + d_P) + (s_L + d_L) (s_P + d_P) K_L K_P}{s_E s_L K_E K_L}.$$

*Proof* Setting all the equations of system (2) to zero, we easily obtain the above results.  $\Box$ 

Lemma 2.1 The set

$$\Delta = \left\{ (x, y, z, w) \in \mathbb{R}^4 | 0 \le x \le K_E, 0 \le y \le K_L, 0 \le z \le K_P, 0 \le w \le \frac{s_P}{f_m} K_P \right\}$$

is positively invariant by the system (2).

*Proof* Let us consider  $(t_0, X_0 = (E_0, L_0, P_0, A_0)) \in \mathbb{R}_+ \times \mathbb{R}^4_+$  and  $([t_0, T_{\max}], X = (E, L, P, A))$  a maximal solution of Cauchy problem associated to (2) with the initial condition  $(t_0, X_0)$ ,  $(T_{\max} \in [t_0, +\infty])$ .

Let us consider  $t_1 \in [t_0, T_{max}]$ . We must show that

- if  $E(t_1) \leq K_E$  then, for all  $t_1 \leq t \leq T_{\text{max}}$ ,  $E(t) \leq K_E$ ,
- if  $L(t_1) \leq K_L$  then, for all  $t_1 \leq t \leq T_{\text{max}}$ ,  $L(t) \leq K_L$ ,
- if  $P(t_1) \leq K_P$  then, for all  $t_1 \leq t \leq T_{\text{max}}$ ,  $P(t) \leq K_P$ ,

- if  $A(t_1) \leq \frac{s_L}{f_m} K_P$  then, for all  $t_1 \leq t \leq T_{\text{max}}$ ,  $A(t) \leq \frac{s_L}{f_m} K_P$ .
- 1. Let us show that, for all  $t \in [t_0, T_{\max}]$ ,  $E(t) \le K_E$ . Assume that there exists  $\epsilon_1 > 0$  such that  $t_1 \le t_1 + \epsilon_1 < T_{\max}$  and  $E(t_1 + \epsilon_1) > K_E$ . We choose  $t_1^* = \inf\{t \ge t_1 | E(t) > K_E\}$ . Since  $E(t_1^*) = K_E$ , a first order limited development of E(t) in the neighborhood of  $t_1^*$  is given by

$$E(t) = K_E + E'(t_1^*)(t - t_1^*) + o(t - t_1^*), \quad t \to t_1^*$$

Besides, using the first equation of system (2), and replacing  $E(t_1^*)$  by  $K_E$ , we obtain

$$E'(t_1^*) = -(s_E + d_E)K_E < 0.$$

So, there exists  $\bar{\epsilon}$  such that, for all  $t_1^* \leq t < t_1^* + \bar{\epsilon}$ ,  $E(t) < K_E$ , this is absurd because of the hypothesis on  $t_1^*$ . We deduce that, for all  $t \in [t_0, T_{\text{max}}]$ ,  $E(t) \leq K_E$ .

2. Now, we want to show that, for all  $t \in [t_0, T_{\text{max}}]$ ,  $L(t) \le K_L$ .

Suppose that there exists  $\epsilon_1$  such that, for all  $t_1 \le t_1 + \epsilon < T_{\text{max}}$  and,  $L(t_1 + \epsilon_1) > K_L$ . Let us assume that  $t_1^* = \inf\{t \ge t_1 | L(t) > K_L\}$ . We have  $L(t_1^*) = K_L$ , so a first order limited development of L(t) about  $t_1^*$  is given by

$$L(t) = K_L + L'(t_1^*)(t - t_1^*) + o(t - t_1^*), \quad t \to t_1^*.$$

From the second equation of system (2), by replacing  $L(t_1^*)$  by  $K_L$  it follows that

 $L'(t_1^*) = -(s_L + d_L)K_L.$ 

This result implies that  $L'(t_1^*) < 0$ . So, there exists  $\bar{\epsilon} > 0$  such that, for all  $t_1^* \le t < t_1^* + \bar{\epsilon}$ ,  $L(t) < K_L$ . This contradicts the hypothesis. Thus, there exists for all  $t \in [t_0, T_{\text{max}}]$ ,  $L(t) \le K_L$ .

3. Let us also show that, for all  $t \in [t_0, T_{\text{max}}]$ ,  $P(t) \le K_P$ .

We suppose that there exists  $\epsilon_1 > 0$  such that  $t_1 \le t_1 + \epsilon_1 < T_{\text{max}}$  and  $P(t_1 + \epsilon_1) > K_P$ . Set  $t_1^* = \inf\{t \ge t_1 | P(t) > K_P\}$ . As  $P(t_1^*) = K_P$ , a first order limited development of P(t) in the neighborhood of  $t_1^*$  is given by

$$P(t) = K_P + P'(t_1^*)(t - t_1^*) + o(t - t_1^*), \quad t \to t_1^*.$$

Furthermore, from the third equation of system (2), we obtain by substituting  $P(t_1^*)$  by  $K_P$ 

 $P'(t_1^*) = -(s_P + d_P)K_P < 0.$ 

Then there exists  $\bar{\epsilon} > 0$  such that, for all  $t_1^* \le t < t_1^* + \bar{\epsilon}$ ,  $P(t) < K_P$ , which is absurd. We deduce that, for all  $t \in [t_0, T_{\text{max}}]$ ,  $P(t) \le K_P$ .

4. Finally, we show that, for all  $t \in [t_0, T_{\max}]$ ,  $A(t) \le \frac{s_P}{f_m} K_P$ . We suppose the existence of  $\epsilon_1 > 0$  such that

$$t_1 \le t_1 + \epsilon_1 < T_{\max}$$
 and  $A(t_1 + \epsilon_1) > \frac{s_P}{f_m} K_P$ .

Considering  $t_1^* = \inf\{t \ge t_1, A(t) > \frac{s_P}{f_m}K_P\}$ , we have  $A(t_1^*) = \frac{s_P}{f_m}K_P$ . Since  $P(t_1^*) = K_P$ , it then follows that

$$A'(t_1^*) = s_P P(t_1^*) - f_m A(t_1^*) = s_P K_P - f_m \times \frac{s_P}{f_m} K_P = 0,$$

namely  $A'(t_1^*) = 0$ . Hence,

$$A''(t_1^*) = s_P P'(t_1^*) - f_m A'(t_1^*)$$
  
=  $s_P P'(t_1^*)$   
=  $-s_P (s_P + d_P) K_P < 0.$ 

A second order limited development of A(t) about  $t_1^*$  yields

$$A(t) = \frac{s_P}{f_m} K_P + A'(t_1^*) (t - t_1^*) + A''(t_1^*) \frac{(t - t_1^*)^2}{2} + o((t - t_1^*)^2), \quad t \to t_1^*.$$

In this case, there exists  $\bar{\epsilon} > 0$  such that, for all  $t_1^* < t \le t_1^* + \bar{\epsilon}, A(t) < \frac{s_P}{f_m} K_P$ . This is absurd.

In conclusion, for all  $t \in [t_0, T_{\max}]$ ,  $A(t) \le \frac{s_P}{f_m} K_P$ .

**Proposition 2.2**  $\Delta$  *is attractive for the system* (2).

*Proof* Let us consider  $(t_0, X_0 = (E_0, L_0, P_0, A_0) \in \mathbb{R}_+ \times \mathbb{R}^4_+ \setminus \Delta$  and  $([t_0, T_{\max}], X = (E, L, P, A))$  a global solution of Cauchy problem associated to (2) with the initial condition  $(t_0, X_0)$ .

Lemma 2.1 shows that  $\Delta$  is invariant. It remains to show that there exists *t* such that  $X(t) \in \Delta$ . We will proceed by showing the contrary to be absurd.

• We suppose that, for all  $t \in [t_0, +\infty[, E(t) > K_E$ . From the first equation of the system (2), we have  $E'(t) = bA(t)(1 - \frac{E(t)}{K_E}) - (s_E + d_E)E(t)$ . Then  $bA(t)(1 - \frac{E(t)}{K_E}) < 0$ , and it follows that  $E'(t) < -(s_E + d_E)K_E$ .

Integrating from  $t_0$  to t, we obtain

$$\int_{t_0}^t E'(t) \, dt \leq - \int_{t_0}^t (s_E + d_E) K_E \, dt, \quad \forall t \geq t_0.$$

Consequently,  $E(t) \leq E(t_0) - (s_E + d_E)K_E(t - t_0), t \leq t_0$ . Posing  $t_1 = t_0 + \frac{E_0(t - K_E)}{(s_E + d_E)K_E}$ , then

$$E(t_1) \le E_0 - (s_E + d_E)K_E \times \left(t_0 + \frac{E_0 - K_E}{(s_E + d_E)K_E} - t_0\right)$$
  
$$\le E_0 - (E_0 - K_E)$$
  
$$\le K_E,$$

which is a contradiction. So, for all  $t > t_1, E(t) \le K_E$ .

• If  $L(t_1) \le K_L$ , then the solution L(t) is defined in  $\Delta$ , which is invariant. If not, suppose that, for any  $t \in [t_1, +\infty[$ , with  $t_1$  previously defined, thus  $L(t) > K_L$ . Then, for all

 $t \in [t_1, +\infty]$ , and thanks to the second equation of the system (2), we have  $L'(t) < -(s_L + d_L)K_L.$ 

By integrating between  $t_1$  and t and using the principle of comparison, we obtain for all  $t \in [t_1, +\infty[: L(t) \le L(t_1) - (s_L + d_L)K_L(t - t_1)]$ . Considering  $t_2 = t_1 + \frac{(L(t_1) - K_L)}{(s_L + d_L)K_L}$ , then

$$\begin{split} L(t_2) &\leq L(t_1) - (s_L + d_L) K_L \left( t_1 + \frac{(L(t_1) - K_L)}{(s_L + d_L) K_L} - t_1 \right) \\ &\leq L(t_1) - (s_L + d_L) K_L \times \frac{(L(t_1) - K_L)}{(s_L + d_L) K_L} \\ &\leq L(t_1) - (L(t_1) - K_L) \\ &\leq K_L. \end{split}$$

Therefore, there exists  $t_2 > t_1$  such that  $L(t_2) \le K_L$ , which is a contradiction. So,  $L(t) < K_L$ 

• If  $P(t) \leq K_P$ , then the solution P(t) is defined in  $\Delta$ , which is invariant. If not, suppose that, for all  $t \in [t_2, +\infty[, P(t) > K_P]$ .

From the third equation of the system (2), we have, for all  $t \in [t_2, +\infty[$ ,  $P'(t) = s_L L(t)(1 - \frac{P(t)}{K_P}) - (s_P + d_P)P(t)$  and as  $(1 - \frac{P(t)}{K_P}) < 0$ , then  $P'(t) < -(s_P + d_P)K_P$ . Integrating from  $t_3$  and t yields

$$P(t) \le P(t_3) - (s_P + d_P)K_P(t - t_3), \quad t \le t_3.$$

Setting  $t_3 = t_2 + \frac{P(t_2) - K_P}{(s_P + d_P)K_P}$ , then

$$P(t_3) \le P(t_2) - (s_P + d_P)K_P \times \left(t_2 + \frac{P(t_2) - K_P}{(s_P + d_P)K_P} - t_2\right)$$
  
$$\le P(t_2) - \left(P(t_2) - K_P\right)$$
  
$$\le K_P.$$

Hence, there exists  $t_3 > t_2$  such that  $P(t_3) \le K_P$ , which is a contradiction. Then,  $\forall t \in [t_3, +\infty[, P(t) \leq K_P].$ 

• If  $A(t) \leq \frac{s_P}{f_m} K_P$ , the solution A(t) is defined in  $\Delta$ , which is invariant. On the other hand, suppose that, for all  $t \in [t_3, +\infty[, A(t) > \frac{s_P}{f_m}K_P]$ . Hence, from the last equation of the system (2), for all  $t \in [t_3, +\infty[, A(t) \le A(t_3) - c(t - t_3)]$ . Considering  $t_4 = t_3 + \frac{A(t_3) - \frac{\delta p}{fm} K_p}{c}$ , it then follows that

$$A(t_4) \leq A(t_3) - c \left( t_3 + \frac{A(t_3) - \frac{s_P}{f_m} K_P}{c} - t_3 \right) \leq \frac{s_P}{f_m} K_P.$$

This is a contradiction.

So, for all  $t > \max(t_1, t_2, t_3, t_4)$ , we have  $(E(t), L(t), P(t), A(t)) \in \Delta$ .

**Corollary 2.1** Let  $(t_0, X_0 = (E_0, L_0, P_0, A_0)) \in \mathbb{R}_+ \times \mathbb{R}_+^4$ . The maximum solution of the problem of Cauchy relative to the system (2) and associated with the initial condition is global. *Proof* Let us consider  $(t, X = (E, L, P, A)) \in \mathbb{R}_+ \times \mathbb{R}^4_+$ , the maximal solution of the Cauchy problem relative to the system (2) and associated with the initial condition  $(t_0, X_0)$ . By Proposition 2.2 and Lemma 2.1, we know that this solution is bounded. Thus, it is global.  $\Box$ 

## 2.2.3 Stability of equilibrium

**Theorem 2.2** The equilibrium  $X_0^* = (0, 0, 0, 0)$  is locally asymptotically stable if and only if r < 1.

*Proof* The local stability of the equilibrium  $X_0^* = (0, 0, 0, 0)$  is given by the Jacobian matrix  $D\mathcal{F}_1(X_0^*)$  of the system evaluated at this point. We have

$$D\mathcal{F}_1(X_0^*) = \begin{pmatrix} -(s_E + d_E) & 0 & 0 & b \\ s_E & -(s_L + d_L) & 0 & 0 \\ 0 & s_L & -(s_P + d_P) & 0 \\ 0 & 0 & s_P & -f_m \end{pmatrix}.$$

The matrix  $D\mathcal{F}_1(X_0^*)$  can be rewritten as  $D\mathcal{F}_1(X_0^*) = M + N$  with M a positive matrix defined by

$$M = \begin{pmatrix} 0 & 0 & 0 & b \\ s_E & 0 & 0 & 0 \\ 0 & s_L & 0 & 0 \\ 0 & 0 & s_P & 0 \end{pmatrix}$$

and N is a diagonal matrix defined as follows:

$$N = \begin{pmatrix} -(s_E + d_E) & 0 & 0 & 0 \\ 0 & -(s_L + d_L) & 0 & 0 \\ 0 & 0 & -(s_P + d_P) & 0 \\ 0 & 0 & 0 & -f_m \end{pmatrix}.$$

Consequently,

$$P = -MN^{-1} = \begin{pmatrix} 0 & 0 & 0 & \frac{b}{f_m} \\ \frac{s_E}{s_E + d_E} & 0 & 0 & 0 \\ 0 & \frac{s_L}{s_L + d_L} & 0 & 0 \\ 0 & 0 & \frac{s_P}{s_P + d_P} & 0 \end{pmatrix}.$$

The characteristic polynomial is given by  $X^4 - r$  and  $\rho(P) = \sqrt[4]{r}$ . From Varga's theorem,  $D\mathcal{F}_1(X_0^*)$  is asymptotically stable if and only if r < 1.

Theorem 2.3 The equilibrium

$$X^* = \left(1 - \frac{1}{r}\right) \left(\frac{K_E}{\chi_E}, \frac{K_L}{\chi_L}, \frac{K_P}{\chi_P}, \frac{s_P}{f_m} \frac{K_P}{\chi_P}\right)$$

is locally asymptotically stable if and only if r > 1.

*Proof* For the proof we evaluate the Jacobian matrix of  $\mathcal{F}_1$  at the endemic equilibrium point  $X^*$ . It can be written as  $D\mathcal{F}_1(X^*) = M + N$  where

$$M = \begin{pmatrix} 0 & 0 & 0 & B \\ C & 0 & 0 & 0 \\ 0 & E & 0 & 0 \\ 0 & 0 & G & 0 \end{pmatrix}$$

with

$$B = b\left(1 - \frac{1}{\chi_E}\left(1 - \frac{1}{r}\right)\right), \qquad C = s_E\left(1 - \frac{1}{\chi_L}\left(1 - \frac{1}{r}\right)\right),$$
$$E = s_L\left(1 - \frac{1}{\chi_P}\left(1 - \frac{1}{r}\right)\right), \qquad G = s_P.$$

Also we have

$$N = \begin{pmatrix} -A & 0 & 0 & 0 \\ 0 & -D & 0 & 0 \\ 0 & 0 & -U & 0 \\ 0 & 0 & 0 & -H \end{pmatrix},$$

where A, D, U and H are, respectively, given by

$$\begin{split} A &= \frac{bs_P K_P}{f_m \chi_P K_E} \left( 1 - \frac{1}{r} \right) + (s_E + d_E), \qquad D = \frac{s_E K_E}{K_L \chi_P} \left( 1 - \frac{1}{r} \right) + (s_L + d_L), \\ U &= \frac{S_L K_L}{K_P \chi_L} \left( 1 - \frac{1}{r} \right) + (s_P + d_P), \qquad H = f_m. \end{split}$$

The matrix *M* is positive if and only if r > 1. On the other hand, the diagonal matrix *N* is invertible and its eigenvalues are all negative if and only if r > 1. Thus,

$$-MN^{-1} = \begin{pmatrix} 0 & 0 & 0 & \alpha_1 \\ \alpha_2 & 0 & 0 & 0 \\ 0 & \alpha_3 & 0 & 0 \\ 0 & 0 & \alpha_4 & 0 \end{pmatrix}$$

with

$$\alpha_{1} = \frac{b(1 - \frac{1}{\chi_{E}}(1 - \frac{1}{r}))}{f_{m}}, \qquad \alpha_{2} = \frac{s_{E}(1 - \frac{1}{\chi_{L}}(1 - \frac{1}{r}))}{\frac{bs_{p}K_{p}}{f_{m}\chi_{P}K_{E}}(1 - \frac{1}{r}) + (s_{E} + d_{E})},$$
$$\alpha_{3} = \frac{s_{L}(1 - \frac{1}{\chi_{P}}(1 - \frac{1}{r}))}{\frac{s_{E}K_{E}}{K_{L}\chi_{P}}(1 - \frac{1}{r}) + (s_{L} + d_{L})}, \qquad \alpha_{4} = \frac{s_{P}}{\frac{s_{L}K_{L}}{K_{P}\chi_{L}}(1 - \frac{1}{r}) + (s_{P} + d_{P})}.$$

The characteristic polynomial of the matrix  $-MN^{-1}$  is  $X^4 - \alpha_1 \alpha_2 \alpha_3 \alpha_4$ , and the spectral radius is given by  $\rho(-MN^{-1}) = \sqrt[4]{\alpha_1 \alpha_2 \alpha_3 \alpha_4}$ .

Since r > 1, the quantity  $\alpha_1 \alpha_2 \alpha_3 \alpha_4$  is less than unity. Consequently,  $\rho(-MN^{-1}) < 1$ , and then the endemic equilibrium is locally asymptotically stable.

**Theorem 2.4** *The trivial equilibrium is globally asymptotically stable if and only if*  $r \le 1$ *.* 

*Proof* Consider the following Lyapunov function [1, 18]:

$$\vartheta(t) = s_L s_P (s_E E(t) + (s_E + d_E)L(t)) + (s_E + d_E)(s_L + d_L) (s_P P(t) + (s_P + d_P)A(t)).$$

By calculating the derivative of  $\vartheta$ , we have

$$\begin{split} \vartheta'(t) &= s_L s_P \Big( s_E E'(t) + (s_E + d_E) L'(t) \Big) + (s_E + d_E) (s_L + d_L) \Big( s_P P'(t) + (s_P + d_P) A'(t) \Big) \\ &= A(t) \Big( b s_E s_L s_P - f_m (s_E + d_E) (s_L + d_L) (s_P + d_P) \Big) \\ &- \Big( s_E s_L s_P \frac{E(t) L(t)}{K_L} + s_L s_P (s_E + d_E) (s_L + d_L) \frac{L(t) P(t)}{K_P} + b s_E s_L s_P \frac{A(t) E(t)}{K_E} \Big) \\ &= \frac{A(t)}{f_m (s_E + d_E) (s_L + d_L) (s_P + d_P)} (r - 1) \\ &- \Big( s_E s_L s_P \frac{E(t) L(t)}{K_L} + s_L s_P (s_E + d_E) (s_L + d_L) \frac{L(t) P(t)}{K_P} + b s_E s_L s_P \frac{A(t) E(t)}{K_E} \Big). \end{split}$$

Since  $r \le 1$ , it then follows that  $\vartheta'(t) \le 0$ . Thanks to LaSalle's invariance principle, the trivial equilibrium,  $X_0^* = (0, 0, 0, 0)$  is globally asymptotically stable.

**Theorem 2.5** *The non-trivial equilibrium is globally asymptotically stable if and only if* r > 1.

*Proof* Suppose that the rate of growth is greater than 1 and  $X^* = (E^*, L^*, P^*, A^*) = (x^*, y^*, z^*, w^*)$ .

Consider the Lyapunov function  $V_1 : \mathbb{R}^4 \to \mathbb{R}$  defined by

$$V_1(x, y, z, w) = \frac{1}{2} (a_1 (x - x^*)^2 + a_2 (y - y^*)^2 + a_3 (z - z^*)^2 + a_4 (w - w^*)^2)$$

with  $a = (a_1, a_2, a_3, a_4)^t \in \mathbb{R}^{*4}_+$  a positive constant vector.

Since r > 1,  $x^*$ ,  $y^*$ ,  $z^*$  and  $w^*$  are also positive.

It is clear that  $V_1(X^*) = 0$  and  $\forall (x, y, z, w) \in \mathbb{R}^4_+ \setminus \{X^*\}, V_1(x, y, z, w) > 0$ . Thus, the function  $V_1$  is well defined and the orbital derivative of  $V_1$  along the solution of the system (2) is

$$V_{1}'(x, y, z, w) = a_{1} \left( x - x^{*} \right) \left( bt \left( 1 - \frac{x}{K_{E}} \right) - (s_{E} + d_{E})x \right)$$
  
+  $a_{2} \left( y - y^{*} \right) \left( s_{E}x \left( 1 - \frac{y}{K_{L}} \right) - (s_{L} + d_{L})y \right)$   
+  $a_{3} \left( z - z^{*} \right) \left( s_{L}y \left( 1 - \frac{z}{K_{P}} \right) - (s_{P} + d_{P})z \right)$   
+  $a_{4} \left( w - w^{*} \right) \left( s_{P}z - f_{m}w \right).$ 

Let us adopt the following notations:

$$\tilde{x} = x - x^*, \qquad \tilde{y} = y - y^*, \qquad \tilde{z} = z - z^*, \qquad \tilde{w} = w - w^*, \qquad \tilde{X} = (\tilde{x}, \tilde{y}, \tilde{z}, \tilde{w})^T.$$

 $\tilde{X}$ 

$$\begin{split} \dot{V}_1(x,y,z,w) &= \tilde{X}^T \begin{pmatrix} -a_1(s_E+d_E) & 0 & 0 & a_1b(1-\frac{x^*}{K_E}) \\ a_2s_E(1-\frac{y^*}{K_L}) & -a_2(s_L+d_L) & 0 & 0 \\ 0 & a_3s_L(1-\frac{z^*}{K_P}) & -a_3(s_P+d_P) & 0 \\ 0 & 0 & a_4s_P & -a_4f_m \end{pmatrix} \\ &- \left(a_1\frac{b}{K_E}x^{*2}w + a_2\frac{s_E}{K_L}y^{*2}x + a_3\frac{s_L}{K_P}z^{*2}y\right). \end{split}$$

Consider  $A_1 = -D + R_1$  with

$$\begin{split} D &= \begin{pmatrix} a_1(s_E + d_E) & 0 & 0 & 0 \\ 0 & a_2(s_L + d_L) & 0 & 0 \\ 0 & 0 & a_3(s_P + d_P) & 0 \\ 0 & 0 & 0 & a_4 f_m \end{pmatrix}; \\ R_1 &= \begin{pmatrix} 0 & 0 & 0 & a_1 b(1 - \frac{x^*}{K_E}) \\ a_2 s_E(1 - \frac{y^*}{K_L}) & 0 & 0 & 0 \\ 0 & a_3 s_L(1 - \frac{z^*}{K_P}) & 0 & 0 \\ 0 & 0 & a_4 s_P & 0 \end{pmatrix}. \end{split}$$

Considering the scalar product of  $\mathbb{R}^4$ , the orbital derivative of the function  $V_1$  can be rewritten in the following form:

$$V_1'(x,y,z,w) = \langle A_1 \tilde{X}, \tilde{X} \rangle - \left( a_1 \frac{b}{K_E} \tilde{x}^2 w + a_2 \frac{s_E}{K_L} \tilde{y}^2 x + a_3 \frac{s_L}{K_P} \tilde{z}^2 y \right).$$

Let us introduce the following symmetric matrix:

$$\begin{split} S_1 &= -D + \frac{1}{2} \Big( R_1^T + R_1 \Big) \\ &= \begin{pmatrix} -a_1(s_E + d_E) & \frac{a_2 s_E}{2} \big( 1 - \frac{y^*}{K_L} \big) & 0 & \frac{a_1 b}{2} \big( 1 - \frac{x^*}{K_E} \big) \\ \frac{a_2 s_E}{2} \big( 1 - \frac{y^*}{K_L} \big) & -a_2 \big( s_L + d_L \big) & \frac{a_3 s_L}{2} \big( 1 - \frac{z^*}{K_P} \big) & 0 \\ 0 & \frac{a_3 s_F}{2} \big( 1 - \frac{z^*}{K_P} \big) & -a_3 \big( s_P + d_P \big) & \frac{a_4 s_P}{2} \\ \frac{a_1 b}{2} \big( 1 - \frac{x^*}{K_E} \big) & 0 & \frac{a_4 s_P}{2} & -a_4 f_m \end{pmatrix} . \end{split}$$

Using the properties of equilibrium states, we get

$$\begin{pmatrix} 1 - \frac{x^*}{K_E} \end{pmatrix} = \frac{s_E + d_E}{b} \times \frac{x^*}{w^*};$$

$$\begin{pmatrix} 1 - \frac{y^*}{K_L} \end{pmatrix} = \frac{s_L + d_L}{s_E} \times \frac{y^*}{x^*};$$

$$\begin{pmatrix} 1 - \frac{z^*}{K_P} \end{pmatrix} = \frac{s_P + d_P}{s_L} \times \frac{z^*}{y^*}.$$

Then

Further, the matrix  $S_1$  becomes

$$S_{1} = \begin{pmatrix} -a_{1}(s_{E} + d_{E}) & \frac{a_{2}}{2}(s_{L} + d_{L})\frac{y^{*}}{x^{*}} & 0 & \frac{a_{1}}{2}(s_{E} + d_{E})\frac{x^{*}}{w^{*}} \\ \frac{a_{2}}{2}(s_{L} + d_{L})\frac{y^{*}}{x^{*}} & -a_{2}(s_{L} + d_{L}) & \frac{a_{3}}{2}(s_{P} + d_{P})\frac{z^{*}}{y^{*}} & 0 \\ 0 & \frac{a_{3}}{2}(s_{P} + d_{P})\frac{z^{*}}{y^{*}} & -a_{3}(s_{P} + d_{P}) & \frac{a_{4}s_{P}}{2} \\ \frac{a_{1}}{2}(s_{E} + d_{E})\frac{x^{*}}{w^{*}} & 0 & \frac{a_{4}s_{P}}{2} & -a_{4}f_{m} \end{pmatrix},$$

and we obtain  $\langle A_1 \tilde{X}, \tilde{X} \rangle = \langle S_1 \tilde{X}, \tilde{X} \rangle$ 

The characteristic polynomial is  $P = X^4 + \gamma_1 X^3 + \gamma_2 X^2 + \gamma_3 X + \gamma_4$  where

$$\begin{split} \gamma_{1} &= a_{1}(s_{E} + d_{E}) + a_{2}(s_{L} + d_{L}) + a_{3}(s_{P} + d_{P}) + a_{4}f_{m}; \\ \gamma_{2} &= a_{1}a_{2}(s_{E} + d_{E})(s_{L} + d_{L}) + a_{3}a_{4}(s_{P} + d_{P})f_{m} + \left(a_{1}(s_{E} + d_{E}) + a_{2}(s_{L} + d_{L})\right) \\ &+ \left(a_{3}(s_{P} + d_{P}) + a_{4}f_{m}\right); \\ \gamma_{3} &= \left(a_{4}a_{3}(s_{P} + d_{P})f_{m}\left(a_{1}(s_{E} + d_{E}) + a_{2}(s_{L} + d_{L})\right) + a_{1}a_{2}(s_{E} + d_{E})(s_{L} + d_{L}) \\ &\times \left(a_{3}(s_{P} + d_{P}) + a_{4}f_{m}\right)\right); \\ \gamma_{4} &= \frac{1}{4}\left(a_{1}a_{3}(s_{E} + d_{E})\left(\frac{x^{*}}{w^{*}}\right)(s_{P} + d_{P})\left(\frac{z^{*}}{y^{*}}\right) - a_{2}a_{4}(s_{L} + d_{L})f_{m}\left(\frac{y^{*}}{x^{*}}\right)\right)^{2} \\ &+ a_{1}a_{2}a_{3}a_{4}f_{m}(s_{E} + d_{E})(s_{L} + d_{L})(s_{P} + d_{P}). \end{split}$$

The global stability of non-trivial equilibrium can be investigated by applying the Routh–Hurwitz criterion on the characteristic polynomial.

The relevant Routh-Hurwitz determinants are

$$\begin{cases} \Delta_1 = \gamma_1 > 0, \\ \Delta_2 = \gamma_1 \gamma_2 - \gamma_3 > 0, \\ \Delta_3 = \gamma_3 \Delta_2 - \gamma_1^2 \gamma_4 > 0, \\ \Delta_4 = \gamma_4 \Delta_3 > 0. \end{cases}$$

It is clear that  $\Delta_1 = \gamma_1 > 0$  and

$$\begin{split} \Delta_2 &= \gamma_1^2 + 2a_1^2a_2(s_E + d_E)^2(s_L + d_L) + a_1a_2^2(s_E + d_E)^2(s_L + d_L)^2 \\ &+ a_3^2a_4(s_P + d_P)^2f_m + a_3a_4^2f_m^2 > 0. \end{split}$$

Let us assume that

$$\begin{aligned} &\alpha = 2a_1^2 a_2 (s_E + d_E)^2 (s_L + d_L) + a_1 a_2^2 (s_E + d_E)^2 (s_L + d_L)^2 + a_3^2 a_4 (s_P + d_P)^2 f_m + a_3 a_4^2 f_m^2, \\ &\Delta_3 = \gamma_3 \Delta_2 - \gamma_1^2 \gamma_4 = \gamma_1^2 (\gamma_3 - \gamma_4) + \alpha \gamma_3 > 0. \end{aligned}$$

Finally, since  $\gamma_4 > 0$ ,  $\Delta_4 = \gamma_4 \Delta_3 > 0$ .

*Remark* 2.1 The above results show that a vector control strategy that brings and maintains the threshold quantity *r*, to a value less than unity will lead to the effective control of

mosquitoes population growth for the community. In other words, the requirement  $r \le 1$  is necessary and sufficient for the effective control of the mosquitoes' population growth [10, 23].

## 3 Malaria transmission dynamics model

## 3.1 Malaria transmission mechanism

Malaria is transmitted to humans by the female of a mosquito of the genus anopheles [24, 28]. There are four species of parasites, which are *Plasmodium falciparum*, *Plasmodium vivax*, *Plasmodium malariae*, and more recently *Plasmodium knowlesi*. However, the most pathogenic is *Plasmodium malariae*, but it remains a rare case. *Plasmodium falciparum* is a frequent case. It causes the most serious illness and is the most widespread in the tropics [27]. Mosquito-to-human malaria transmission occurs when sporozoites from the salivary gland of the mosquito are injected into the skin during blood-feeding. Parasites then pass to the liver where they replicate, each sporozoite yielding many thousands of merozoites which go on to cause patent infection. The biology of the four species of *Plasmodium* is generally similar and consists of two distinct phases: a sexual stage at the mosquito host and an asexual stage at the human host. The asexual phase consists of at least three forms: sporozoites, merozoites, and trophozoites. During the asexual stage, some of the parasites become gametocytes and then when a mosquito bites an infected human, it ingests the gametocytes. Hence, the parasite continues its development and invades the salivary glands of the mosquito ending the cycle [13, 27].

## 3.2 Model formulation of malaria transmission

In this section, we give a brief description of the different stages of our model of malaria parasite transmission. In order to derive our model, we divide the human population into two major types. The first type, called non-immune, is divided into three sub-classes, and the second type, called semi-immune, is divided in four classes. Tables 1-6 give a description of all these classes.

The total human population at each instant t is given by

$$N_h(t) = S_e(t) + E_e(t) + I_e(t) + S_a(t) + E_a(t) + I_a(t) + R_a(t).$$
(5)

Table 1	Parameters	for	human	hosts

Notation	Description
S <sub>e</sub> S <sub>a</sub> E <sub>e</sub> E <sub>a</sub>	class of non-immune susceptible class of semi-immune susceptible class of non-immune latent class of semi-immune infectious class of non-immune infectious
r <sub>a</sub> Ra	class of immune semi-immune

<b>Table 2</b> Parameters for vector
--------------------------------------

Notation	Description
S <sub>m</sub> E <sub>m</sub>	class of susceptible mosquitoes class of latent mosquitoes
Im	class of infectious mosquitoes

#### Table 3 Parameters common to non-immune and semi-immune

Parameters	Biological description
$\Lambda_h$	constant recruitment rate of human (it also includes births)
р	probability for a new recruit to be non-immune
1 – <i>p</i>	probability for a new recruit to be semi-immune

#### Table 4 Parameters for semi-immune hosts

Parameters	Biological description
$v_a$	rate of passage from latent semi-immune to infectious
$\alpha_a$	rate of passage infectious semi-immune to immune
$oldsymbol{eta}_a$	rate of loss of immune immunity

Tabl	e 5	Contact	parameters	between	non-immune,	semi-imm	une and	mosq	uito
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Parameters	Biological description
n <sub>a</sub>	average number of bites per mosquito per unit of time
C <sub>me</sub>	probability that an infectious mosquito bite on a susceptible non-immune transfers the infection to the non-immune
C <sub>ma</sub>	probability that an infectious mosquito bite on a susceptible semi-immune transfers the infection to the semi-immune
C <sub>em</sub>	probability that a bite from a susceptible mosquito on an infectious non-immune transfers the infection to the mosquito
C <sub>am</sub>	probability that a bite from a susceptible mosquito on an infectious semi-immune transfers the infection to the mosquito
С <sub>ат</sub>	probability that a bite from a susceptible mosquito on an infectious immune transfers the infection to the mosquito

 Table 6
 Parameters for the non-immune hosts

Parameters	Biological description
$\nu_e$	rate of passage from latent non-immune to infectious
$\alpha_e$	rate of passage infectious non-immune to immune
$\gamma_e$	mortality rate due to malaria on non-immune
γα	mortality rate due to malaria on semi-immune

The total vector population at each instant *t* is given by

$$A(t) = S_m(t) + E_m(t) + I_m(t).$$

We make the following useful assumptions.

- (H3): We assume that an immigrant is either non-immune or semi-immune.
- (H4): We assume that the only mode of transmission is the mosquito bites.
- (H5): It is assumed that an individual who arrives newly in our study area has a probability p of being non-immune and a probability 1 p of being semi-immune.
- (H6): We assume that all recruits are susceptible.
- (H7): We assume that the natural mortality rate  $f_h$  (resp.  $f_m$ ) is constant.
- (H8): The probabilities  $c_{me}$ ,  $c_{ma}$ ,  $c_{em}$ ,  $c_{am}$ ,  $\tilde{c}_{am}$ , the parameters  $v_e$ ,  $v_a$ ,  $v_m$ ,  $\alpha_e$ ,  $\alpha_a$ ,  $\beta_a$  and  $n_a$  are and the induced mortality rates  $\gamma_e$  and  $\gamma_a$  are non-negative.

The forces of infections from mosquitoes to non-immune and semi-immune are, respectively, defined by

$$k_e = c_{me} n_a \frac{I_m}{N_h},$$

(6)

(7)



$$k_a = c_{ma} n_a \frac{I_m}{N_h}$$

and the force of infection from human to mosquitoes is

$$k_m = c_{am} n_a \frac{I_a}{N_h} + c_{em} n_a \frac{I_e}{N_h} + \tilde{c}_{am} n_a \frac{R_a}{N_h}.$$

(H9): Let us assume that  $0 < v_e \le k_e$ ,  $0 < v_a \le k_a$  and  $0 < v_m \le k_m$ . Taking into account all these above hypotheses, the inter-host dynamics has been illustrated as in [13].

Thus, the overall dynamics of the spread of the disease is reflected in a diagram. From Fig. 2, by making the balance in each compartment, we obtain the following system of ordinary differential equations:

$$\begin{cases} E'(t) = bA(t)(1 - \frac{E(t)}{K_E}) - (s_E + d_E)E(t), \\ L'(t) = s_E E(t)(1 - \frac{L(t)}{K_L}) - (s_L + d_L)L(t), \\ P'(t) = s_L L(t)(1 - \frac{P(t)}{K_P}) - (s_P + d_P)P(t), \end{cases}$$

$$\begin{cases} S'_e(t) = p\Lambda_h - (f_h + k_e)S_e(t), \\ E'_e(t) = k_eS_e(t) - (f_h + v_e)E_e(t), \\ I'_e(t) = v_eE_e(t) - (f_h + \gamma_e + \alpha_e)I_e(t), \\ S'_a(t) = (1 - p)\Lambda_h + \beta_a R_a(t) - (f_h + k_a)S_a(t), \\ E'_a(t) = k_a S_a(t) - (f_h + v_a)E_a(t), \\ I'_a(t) = v_a E_a(t) - (f_h + \gamma_a + \alpha_a)I_a(t), \\ R'_a(t) = \alpha_e I_e(t) + \alpha_a I_a(t) - (f_h + \beta_a)R_a(t), \\ S'_m(t) = s_P P(t) - (f_m + k_m)S_m(t), \\ E'_m(t) = k_m S_m(t) - (f_m + v_m)E_m(t), \\ I'_m(t) = v_m E_m(t) - f_m I_m(t). \end{cases}$$
(S1)

In system (7), the subsystem (S1) represents the vectors population growth global dynamics and subsystem (S2) represents the virus transmission. Indeed, (S1) is coupled with (S2) through the variable P(t). However, to analyze the system (7) in a decoupled form, we can use the principle of limiting system [16, 36].

From Eq. (6), we obtain

$$S_m(t) = A(t) - E_m(t) - I_m(t),$$
(8)

and then, replacing the expression of  $S_m(t)$  given by Eq. (8) into the system (7), it is easy to see that the system (7) is equivalent to the following one:

$$\begin{cases} E'(t) = bA(t)(1 - \frac{E(t)}{K_E}) - (s_E + d_E)E(t), \\ L'(t) = s_E E(t)(1 - \frac{I(t)}{K_L}) - (s_L + d_L)L(t), \\ P'(t) = s_L L(t)(1 - \frac{P(t)}{K_P}) - (s_P + d_P)P(t), \\ A'(t) = s_P P(t) - f_m A(t), \end{cases}$$

$$\begin{cases} S'_e(t) = p\Lambda_h - (f_h + k_e)S_e(t), \\ E'_e(t) = k_e S_e(t) - (f_h + v_e)E_e(t), \\ I'_e(t) = v_e E_e(t) - (f_h + \gamma_e + \alpha_e)I_e(t), \\ E'_a(t) = (1 - p)\Lambda_h + \beta_a R_a(t) - (f_h + k_a)S_a(t), \\ E'_a(t) = k_a S_a(t) - (f_h + v_a)E_a(t), \\ I'_a(t) = v_a E_a(t) - (f_h + \gamma_a + \alpha_a)I_a(t), \\ R'_a(t) = \alpha_e I_e(t) + \alpha_a I_a(t) - (f_h + \beta_a)R_a(t), \\ E'_m(t) = k_m (A(t) - E_m(t) - I_m(t)) - (f_m + v_m)E_m(t), \\ I'_m(t) = v_m E_m(t) - f_m I_m(t). \end{cases}$$
(S1)

At any time  $t \ge 0$ , the total size of the humans population and adult mosquitoes population are, respectively, given by the following equations:

$$N'_{h}(t) = \Lambda_{h} - f_{h}N_{h} - \gamma_{e}I_{e} - \gamma_{a}I_{a}, \qquad (10)$$

$$A'(t) = s_P P - f_m A(t).$$
(11)

*Remark* 3.1 The previous results indicate that the mosquito population will die out if the vector threshold r is less than or equal to unity, while the mosquito population will eventually stabilize at a positive equilibrium ( $E^*$ ,  $L^*$ ,  $P^*$ ,  $A^*$ ) if the vector threshold r is greater than unity.

From the system (9), we obtain the following limit system:

$$\begin{cases} S'_{e}(t) = p\Lambda_{h} - (f_{h} + k_{e})S_{e}(t), \\ E'_{e}(t) = k_{e}S_{e}(t) - (f_{h} + \nu_{e})E_{e}(t), \\ I'_{e}(t) = \nu_{e}E_{e}(t) - (f_{h} + \gamma_{e} + \alpha_{e})I_{e}(t), \\ \\ S'_{a}(t) = (1 - p)\Lambda_{h} + \beta_{a}R_{a}(t) - (f_{h} + k_{a})S_{a}(t), \\ E'_{a}(t) = k_{a}S_{a}(t) - (f_{h} + \nu_{a})E_{a}(t), \\ I_{a}(t) = \nu_{a}E_{a}(t) - (f_{h} + \gamma_{a} + \alpha_{a})I_{a}(t), \\ R'_{a}(t) = \alpha_{e}I_{e}(t) + \alpha_{a}I_{a}(t) - (f_{h} + \beta_{a})R_{a}(t), \\ \\ \\ E'_{m}(t) = k_{m}(A^{*} - E_{m}(t) - I_{m}(t)) - (f_{m} + \nu_{m})E_{m}(t), \\ I'_{m}(t) = \nu_{m}E_{m}(t) - f_{m}I_{m}(t). \end{cases}$$

$$(12)$$

## 3.3 Mathematical analysis of malaria transmission model

In this part of the paper, we focus on the study of the system (12) under the influence of the mosquito growth rate [11, 13, 15, 23].

Note that the system (12) can be represented as follows:

$$X'(t) = \mathcal{F}_2(X(t))$$

where

$$X(t) = (S_e, E_e, I_e, S_a, E_a, I_a, R_a, E_m, I_m)^T$$

and

$$\mathcal{F}_{2}(X) = \begin{pmatrix} p\Lambda_{h} - (f_{h} + k_{e})S_{e} \\ k_{e}S_{e} - (f_{h} + \nu_{e})E_{e} \\ \nu_{e}E_{e} - (f_{h} + \gamma_{e} + \alpha_{e})I_{e} \\ (1 - p)\Lambda_{h} + \beta_{a}R_{a} - (f_{h} + k_{a})S_{a} \\ k_{a}S_{a} - (f_{h} + \nu_{a})E_{a} \\ \nu_{a}E_{a} - (f_{h} + \gamma_{a} + \alpha_{a})I_{a} \\ \alpha_{e}I_{e} + \alpha_{a}I_{a} - (f_{h} + \beta_{a})R_{a} \\ k_{m}(A^{*} - E_{m} - I_{m}) - (f_{m} + \nu_{m})E_{m} \\ \nu_{m}E_{m} - f_{m}I_{m} \end{pmatrix}.$$

## 3.3.1 Existence and positivity of solutions

**Lemma 3.1** For any initial conditions, the system (12) has a unique positive solution for all  $t \ge 0$ . Further, the domain  $\Delta' = \Omega_h \times \Omega_m \subset \mathbb{R}^9_+$  where

$$\Omega_h = \left\{ (S_e, E_e, I_e, S_a, E_a, I_a, R_a) | 0 \le N_h \le \frac{\Lambda_h}{f_h} \right\}$$

and

$$\Omega_m = \left\{ (E_m, I_m) | 0 \le E_m + I_m \le \frac{s_P}{f_m} K_P \right\}$$

is positively invariant and attracts all the positive orbits of  $\mathbb{R}_+$ .

*Proof* Since  $\mathcal{F}_2$  is  $\mathcal{C}^1$ , it is locally Lipschitzian on  $\mathbb{R}^9$ , we deduce the existence and uniqueness of the solution to the Cauchy problem associated with the subsystem relative to the initial condition  $(t_0, X_0) \in \mathbb{R} \times \mathbb{R}^9$ . Since  $\mathcal{F}_2$  is  $\mathcal{C}^\infty$ , we deduce that the solution is also  $\mathcal{C}^\infty$ . Now, assuming that there is no disease induced death rate, (10) becomes

$$N_h'(t) = \Lambda_h - f_h N_h. \tag{13}$$

Let us assume that  $N'_h(t) \le 0$  and  $N'_m(t) \le 0$ .

It follows that  $N_h \leq \frac{\Lambda_h}{f_h}$ ,  $A \leq \frac{s_P}{f_m} K_P$ .

Then, as in [6, 19], the following inequalities hold:

$$0 \le N_h \le \frac{\Lambda_h}{f_h}$$
,  $0 \le A \le \frac{s_P}{f_m} K_P$ .

Therefore, Eqs. (11) and (13), respectively, become

$$N_h'(t) \le \Lambda_h - f_h N_h$$

and

$$A'(t) \le \Lambda_m - f_m A(t).$$

Using the variation of constant method, between t and  $t_0$ , we have the following solutions:

$$\begin{split} N_h(t) &= \frac{\Lambda_h}{f_h} + \left(N_h^0 - \frac{\Lambda_h}{f_h}\right) e^{-f_h(t-t_0)},\\ A(t) &= \frac{S_P K_P}{f_m} + \left(A_0 - \frac{\Lambda_m}{f_m}\right) e^{-f_m(t-t_0)}. \end{split}$$

From the theorem of comparison, it follows that

$$N_h(t) \leq rac{\Lambda_h}{f_h} + \left(N_h^0 - rac{\Lambda_h}{f_h}
ight) e^{-f_h(t-t_0)}$$

and

$$A(t) \leq \frac{S_P K_P}{f_m} + \left(A_0 - \frac{\Lambda_m}{f_m}\right) e^{-f_m(t-t_0)}.$$

So, the total size of the humans population  $N_h(t) \rightarrow \frac{\Lambda_h}{f_h}$  as  $t \rightarrow \infty$ . Similarly, the total size of the mosquitoes population  $A \rightarrow \frac{S_P K_P}{f_m}$  as  $t \rightarrow \infty$ .

It implies that the set  $\Delta'$  is bounded and we deduce the global existence of the solutions in  $[0, +\infty[$ .

However, assuming that  $N_h(t) > \frac{\Lambda_h}{f_h}$  (respectively,  $A(t) > \frac{s_P}{f_m}K_P$ ), we obtain  $N'_h(t) < \Lambda_h - f_h \times \frac{\Lambda_h}{f_h}$ , namely,  $N'_h(t) < 0$ .

In this case, the two hosts size would be decreasing. Since the domain  $\Delta'$  is compact, all the solutions remain there.

## 3.3.2 Basic reproduction number

By linearizing the system in the neighborhood of the trivial equilibrium point  $DFE_0$ , we obtain the following linear differential system: X'(t) = BX(t) where *B* denotes the Jacobian matrix of the function  $\mathcal{F}_1$  at the equilibrium point  $DFE_0$ , and it is defined as follows:

$$B = \begin{pmatrix} B_{11} & B_{12} \\ B_{21} & B_{22} \end{pmatrix},$$

where the matrices  $B_{11}$ ,  $B_{12}$ ,  $B_{21}$  and  $B_{22}$  are, respectively, given by

$$B_{11} = \begin{pmatrix} -B^* & 0 & 0 & 0 & 0 & 0 & c_{me}n_a \frac{S_e^*}{N_h^*} \\ v_e & -C^* & 0 & 0 & 0 & 0 & 0 \\ 0 & 0 & -D^* & 0 & 0 & 0 & c_{ma}n_a \frac{S_a^*}{N_h^*} \\ 0 & 0 & v_a & -E^* & 0 & 0 & 0 \\ 0 & \alpha_e & 0 & \alpha_a & -G^* & 0 & 0 \\ 0 & c_{em}n_a \frac{A^*}{N_h^*} & 0 & c_{am}n_a \frac{A^*}{N_h^*} & \tilde{c}_{am}n_a \frac{A^*}{N_h^*} & -H^* & 0 \\ 0 & 0 & 0 & 0 & 0 & v_m & -f_m \end{pmatrix}$$

with

The sub-matrix  $B_{11}$  is called *transmission matrix* and it is Metzler stable. It can be decomposed as  $B_{11} = F + V$  where

	/ D*	0	0	0	0	0	()	
	-B	0	0	0	0	0	0	
	ve	$-C^*$	0	0	0	0	0	
	0	0	$-D^*$	0	0	0	0	
<i>V</i> =	0	0	$v_a$	$-E^*$	0	0	0	
	0	$\alpha_e$	0	$\alpha_a$	$-G^*$	0	0	
	0	0	0	0	0	$-H^*$	0	
	0	0	0	0	0	$v_m$	$-f_m$	

The inverse of the matrix V is

$$V^{-1} = \begin{pmatrix} -\frac{1}{B^*} & 0 & 0 & 0 & 0 & 0 & 0 \\ -\frac{\nu_c}{B^*C^*} & -\frac{1}{C^*} & 0 & 0 & 0 & 0 & 0 \\ 0 & 0 & -\frac{1}{D^*} & 0 & 0 & 0 & 0 \\ 0 & 0 & -\frac{\nu_a}{D^*E^*} & -\frac{1}{E^*} & 0 & 0 & 0 \\ 0 & 0 & -\frac{\alpha_a\nu_a}{D^*E^*G^*} & -\frac{\alpha_a}{E^*G^*} & -\frac{1}{G^*} & 0 & 0 \\ 0 & 0 & 0 & 0 & 0 & -\frac{1}{H^*} & 0 \\ 0 & 0 & 0 & 0 & 0 & 0 & -\frac{\nu_m}{f_mH^*} & -\frac{1}{f_m} \end{pmatrix}.$$

**Definition 3.1** The matrix

$$-FV^{-1} = \begin{pmatrix} 0 & 0 & 0 & 0 & 0 & 0 & K_{17} \\ 0 & 0 & 0 & 0 & 0 & 0 & K_{27} \\ 0 & 0 & 0 & 0 & 0 & 0 & K_{37} \\ 0 & 0 & 0 & 0 & 0 & 0 & K_{47} \\ 0 & 0 & 0 & 0 & 0 & 0 & K_{57} \\ 0 & K_{62} & 0 & K_{64} & K_{65} & 0 & 0 \\ 0 & K_{72} & 0 & K_{74} & K_{75} & 0 & 0 \end{pmatrix}$$

is called the next generation matrix. The non-zero coefficients are given by

$$\begin{split} K_{17} &= \frac{1}{B^*} \times c_{me} n_a \frac{S_e^*}{N_h^*}, \qquad K_{27} = \frac{\nu_e}{B^* C^*} \times c_{me} n_a \frac{S_e^*}{N_h^*}, \qquad K_{37} = \frac{1}{D^*} \times c_{ma} n_a \frac{S_a^*}{N_h^*}, \\ K_{47} &= \frac{\nu_a}{D^* E^*} \times c_{ma} n_a \frac{S_a^*}{N_h^*}, \qquad K_{57} = \frac{\alpha_a \nu_a}{D^* E^* G^*} \times c_{ma} n_a \frac{S_a^*}{N_h^*}, \qquad K_{62} = \frac{1}{H^*} \times c_{em} n_a \frac{A^*}{N_h^*}, \\ K_{64} &= \frac{1}{H^*} \times c_{am} n_a \frac{A^*}{N_h^*}, \qquad K_{65} = \frac{1}{H^*} \times \tilde{c}_{am} n_a \frac{A^*}{N_h^*}, \qquad K_{72} = \frac{\nu_m}{H^*} \times \frac{1}{f_m} \times c_{em} n_a \frac{A^*}{N_h^*}, \\ K_{74} &= \frac{\nu_m}{H^*} \times \frac{1}{f_m} \times c_{am} n_a \frac{A^*}{N_h^*}, \qquad K_{75} = \frac{\nu_m}{H^*} \times \frac{1}{f_m} \times \tilde{c}_{am} n_a \frac{A^*}{N_h^*}. \end{split}$$

**Proposition 3.1** The basic reproduction number is

$$\mathcal{R}_0 = \rho\left(-FV^{-1}\right) = \sqrt{K_{27}K_{72} + K_{47}K_{74} + K_{57}K_{75}}.$$
(14)

*Proof* According to the mathematical sense, the basic reproduction number is the spectral radius of next generation matrix. For this purpose, the above expression of  $\mathcal{R}_0$  is obtained just by using this definition and making some calculations.

and

**Proposition 3.2** *The system* (12) *has a unique disease-free equilibrium given by* 

$$DFE_0 = \left(\frac{p}{f_h}\Lambda_h, 0, 0, \frac{1-p}{f_h}\Lambda_h, 0, 0, 0, 0, 0\right).$$

*Proof* When we consider the case where there are any mosquitoes and infected humans, we have  $E_m^* = I_m^* = 0$  and  $E_e^* = I_e^* = E_a^* = I_a^* = R_a^* = 0$ ; then the trivial equilibrium is  $X_0^* = (S_e^*, 0, 0, S_a^*, 0, 0, 0, 0, 0, 0)$ . Furthermore, without infected classes, the forces of infection are equal to zero, and then  $S_e^*$  and  $S_a^*$  are obtained solving the following equations:

$$\begin{split} S'_{e}(t) &= 0 \quad \Leftrightarrow \quad p \Lambda_{h} - f_{h} S^{*}_{e} = 0 \\ &\Leftrightarrow \quad S^{*}_{e} = \frac{p}{f_{h}} \Lambda_{h}, \\ S'_{a}(t) &= 0 \quad \Leftrightarrow \quad (1 - p)\Lambda_{h} - f_{h} S^{*}_{a} = 0 \\ &\Leftrightarrow \quad S^{*}_{a} = \frac{1 - p}{f_{h}} \Lambda_{h}, \\ N^{*}_{h} &= S^{*}_{e} + S^{*}_{a} = \frac{\Lambda_{h}}{f_{h}}. \end{split}$$

Finally,  $DFE_0 = (\frac{p}{f_h} \Lambda_h, 0, 0, \frac{1-p}{f_h} \Lambda_h, 0, 0, 0, 0, 0).$ 

#### 3.3.3 Stability of equilibrium states

**Theorem 3.1** The disease-free equilibrium  $DFE_0$  is locally asymptotically stable if  $\mathcal{R}_0$  is less than unity and instable if  $\mathcal{R}_0$  is greater than unity.

*Proof* Indeed, the disease-free equilibrium  $DFE_0$  is stable when the spectral radius of the next generation matrix is less than unity. As  $\rho(-FV^-1) = \mathcal{R}_0$ , we deduce the result.  $\Box$ 

Let us consider the following region:

$$\Delta^* = \left\{ (S_e, E_e, I_e, S_a, E_a, I_a, R_a, E_m, I_m) \in \mathbb{R}^9_+ | 0 < S_e \le S^*_e, \dots, 0 < I_m \le I^*_m \right\}.$$
(15)

It is clear that  $\Delta^* \subset \Delta$ . Since  $\Delta$  is positively invariant, we deduce that  $\Delta^*$  is positively invariant.

**Theorem 3.2** If  $\mathcal{R}_0 < 1$ , then the DFE  $X_1^*$  is globally asymptotically stable in  $\Delta^*$ .

*Proof* Let us consider the following Lyapunov function:

$$\mathcal{L}(t) = f_1 E_e + f_2 I_e + f_3 E_a + f_4 I_a + f_5 R_a + f_6 E_m + f_7 I_m,$$

where

$$\chi = \frac{k_e}{B^*} \times \frac{\nu_e}{C^*} \times \frac{k_a}{D^*} \times \frac{\nu_a}{E^*} \times \frac{\alpha_e + \alpha_a}{G^*} \times \frac{k_m}{H^* + k_m} \times \frac{\nu_m}{f_m},$$
  
$$f_1 = \chi \mathcal{R}_0, \qquad f_2 = \frac{B^*}{k_e} f_1, \qquad f_3 = \frac{C^*}{\nu_e} f_2, \qquad f_4 = \frac{D^*}{k_a} f_3,$$

$$f_5 = 0,$$
  $f_6 = \frac{E^*}{v_a} \times \frac{G^*}{\alpha_e + \alpha_a} f_4,$   $f_7 = \frac{f_m}{v_m} f_6,$ 

with Lyapunov derivative given by

$$\begin{aligned} \mathcal{L}'(t) &= f_1 E'_e + f_2 I'_e + f_3 E'_a + f_4 I'_a + f_5 R'_a + f_6 E'_m + f_7 I'_m, \\ \mathcal{L}'(t) &= f_1 (k_e S_e - B^* E_e) + f_2 (v_e E_e - C^* I_e) + f_3 (k_a S_a - D^* E_a) \\ &+ f_4 (v_a E_a - E^* I_a) + f_6 (k_m A^* - k_m I_m - (H^* + k_m) E_m) \\ &+ f_7 (v_m E_m - f_m I_m) \\ &= k_e \chi S_e (\mathcal{R}_0 - 1) + (f_2 v_e E_e - k_e f_2 E_e) + (f_4 v_a E_a - k_a f_4 E_a) \\ &+ \left(\frac{f_m}{v_m} f_6 E_m - f_6 (H^* + k_m) E_m\right) + (k_e \chi S_e - f_2 C^* I_e) \\ &+ (f_3 k_a S_a - f_4 E^* I_a) + (f_6 k_m A^* - f_6 k_m I_m - f_7 f_m I_m) \\ &\leq k_e \chi S_e (\mathcal{R}_0 - 1) + \mathcal{R}_0 \chi_1 E_e \left(\frac{v_e}{k_e} - 1\right) + \mathcal{R}_0 \chi_2 E_a \left(\frac{v_a}{k_a} - 1\right) \\ &+ \mathcal{R}_0 \chi_3 E_m \left(\frac{f_m}{v_m (H^* + k_m)} - 1\right) + 2 \left(\frac{k_e^2}{\mathcal{R}_0 B^* C^*} - 1\right) C^* f_2 \max(S_e^*, I_e^*) \\ &+ 2 \left(\frac{k_a^2}{D^*} - 1\right) f_4 \max(S_a^*, I_a^*) - 2 f_7 f_m \max(A^*, I_m^*) \\ &\leq k_e \chi S_e (\mathcal{R}_0 - 1) + \chi_1 \mathcal{R}_0 E_e \left(\frac{v_e}{k_e} - 1\right) + \chi_2 \mathcal{R}_0 E_a \left(\frac{v_a}{k_a} - 1\right) \\ &+ \mathcal{R}_0 \chi_3 E_m \left(\frac{f_m}{v_m (H^* + k_m)} - 1\right), \end{aligned}$$

where

$$\begin{split} \chi_1 &= k_e \frac{v_e}{C^*} \times \frac{k_a}{D^*} \times \frac{v_a}{E^*} \times \frac{\alpha_e + \alpha_a}{G^*} \times \frac{k_m}{H^* + k_m} \times \frac{v_m}{f_m}, \\ \chi_2 &= k_a \frac{v_a}{E^*} \times \frac{\alpha_e + \alpha_a}{G^*} \times \frac{k_m}{H^* + k_m} \times \frac{v_m}{f_m}, \\ \chi_3 &= k_m \frac{v_m}{f_m}. \end{split}$$

So, if  $\mathcal{R}_0 < 1$ , then  $\mathcal{L}'(t) \leq 0$ .

## **Theorem 3.3** If $\mathcal{R}_0 > 1$ then the system (12) has a unique endemic equilibrium.

*Proof* Let be  $X^{**}$  the non-trivial equilibrium; the components  $S_e^{**}$ ,  $E_e^{**}$ ,  $I_e^{**}$ ,  $S_a^{**}$ ,  $E_a^{**}$ ,  $I_a^{**}$ ,  $R_a^{**}$ ,  $E_m^{**}$ ,  $I_m^{**}$  of  $X^{**}$  are obtained setting all the equations of the system (12) to zero. Hence,

$$\begin{split} S_e^{**} &= \frac{p}{f_h + k_e^{**}} \Lambda_h, \qquad E_e^{**} = \frac{pk_e^{**}}{(f_h + k_e)B^*} \Lambda_h, \qquad I_e^{**} = \frac{pk_e^{**}v_e}{(f_h + k_e^{**})B^*C^*} \Lambda_h, \\ S_a^{**} &= \frac{(pk_e^{**}\alpha_e v_e - (1-p)(f_h + k_e^{**})B^*C^*G^*)D^*E^*}{((f_h + k_a^{**})D^*E^*G^* - k_a^{**}\alpha_a v_a \beta_a)(f_h + k_e^{**})B^*C^*}, \\ E_a^{**} &= \frac{(pk_e^{**}\alpha_e v_e - (1-p)(f_h + k_e^{**})B^*C^*G^*)k_a^{**}E^*}{((f_h + k_a^{**})D^*E^*G^* - k_a^{**}\alpha_a v_a \beta_a)(f_h + k_e^{**})B^*A^*}, \end{split}$$

$$\begin{split} I_a^{**} &= \frac{(pk_e^{**}\alpha_e v_e - (1-p)(f_h + k_e^{**})B^*C^*G^*)k_a^{**}v_a}{((f_h + k_a^{**})D^*E^*G^* - k_a^{**}\alpha_a v_a \beta_a)(f_h + k_e^{**})B^*C^*}, \\ R_a^{**} &= \frac{(pk_e^{**}\alpha_e v_e - (1-p)(f_h + k_e^{**})B^*C^*G^*)(f_h + k_a^{**})D^*E^*}{((f_h + k_a^{**})D^*E^*G^* - k_a^{**}\alpha_a v_a \beta_a)(f_h + k_e^{**})\beta_a B^*C^*} - \frac{1-p}{\beta_a}\Lambda_h, \\ E_m^{**} &= \left(1 - \frac{1}{r}\right)\frac{k_m^{**}s_P K_P}{(f_m + 1)(f_m + v_m)\chi_P}, \qquad I_m^{**} = \left(1 - \frac{1}{r}\right)\frac{k_m^{**}v_m s_P K_P}{f_m(f_m + 1)(f_m + v_m)\chi_P H^*}. \quad \Box$$

**Theorem 3.4** If  $\mathcal{R}_0 > 1$ , then the system (12) has a unique endemic equilibrium which is globally asymptotically stable in the following set:

$$\Delta^{**} = \left\{ (S_e, E_e, I_e, S_a, E_a, I_a, R_a, E_m, I_m) \in \mathbb{R}^9_+ | 0 < S_e \le S_e^{**}, \dots, 0 < I_m \le I_m^{**} \right\}.$$

*Proof* Since  $\mathcal{R}_0 > 1$ , the endemic equilibrium exists. Now, let us consider the following Lyapunov function [3]:

$$\mathcal{V}(t) = \mathcal{V}_1(t) + \mathcal{V}_2(t),$$

where

$$\mathcal{V}_{1}(t) = \frac{1}{2} \left( V_{se}(t) + V_{ee}(t) + V_{ie}(t) + V_{sa}(t) + V_{ea}(t) + V_{ia}(t) + V_{ra}(t) \right)^{2}$$

and

$$\mathcal{V}_2(t) = \frac{1}{2} \left( V_{em}(t) + V_{im}(t) \right)^2$$

with

$$\begin{aligned} V_{se}(t) &= S_e(t) - S_e^{**}, & V_{ee}(t) = E_e(t) - E_e^{**}, & V_{ie}(t) = I_e(t) - I_e^{**}, \\ V_{sa}(t) &= S_a(t) - S_a^{**}, & V_{ea}(t) = E_a(t) - E_a^{**}, & V_{ia}(t) = I_a(t) - I_a^{**}, \\ V_{ra}(t) &= R_a(t) - R_a^{**}, & V_{em}(t) = E_m(t) - E_m^{**}, & V_{im}(t) = I_m(t) - I_m^{**}. \end{aligned}$$

The Lyapunov function constructed above guarantees that it attains the minimum value at the endemic equilibrium  $(S_e^{**}, E_e^{**}, I_e^{**}, S_a^{**}, E_a^{**}, I_a^{**}, R_a^{**}, E_m^{**}, I_m^{**})$ .

The Lyapunov derivative of this function is given by

$$\mathcal{V}'(t) = \mathcal{V}'_1(t) + \mathcal{V}'_2(t)$$

with

$$\mathcal{V}'_{1}(t) = \left(V_{se}(t) + V_{ee}(t) + V_{ie}(t) + V_{sa}(t) + V_{ea}(t) + V_{ia}(t) + V_{ra}(t)\right) \times N'_{h}(t)$$

and

$$\mathcal{V}_2'(t) = \left(V_{em}(t) + V_{im}(t)\right) \times \left(E_m'(t) + I_m'(t)\right).$$

Consequently,

$$\begin{aligned} \mathcal{V}'(t) &= \left( V_{se}(t) + V_{ee}(t) + V_{ie}(t) + V_{sa}(t) + V_{ea}(t) + V_{ia}(t) + V_{ra}(t) \right) \times N_h'(t) \\ &+ \left( V_{em}(t) + V_{im}(t) \right) \times \left( E_m'(t) + I_m'(t) \right). \end{aligned}$$

It is easy to see that  $\mathcal{V}'(t) = 0$  if and only if  $S_e = S_e^{**}$ ,  $E_e = E_e^{**}$ ,  $I_e = I_e^{**}$ ,  $S_a = S_a^{**}$ ,  $E_a = E_a^{**}$ ,  $R_a = R_a^{**}$ ,  $I_a = I_a^{**}$ ,  $E_m = E_m^{**}$ ,  $I_m = I_m^{**}$ .

Now, let us show that  $\mathcal{V}'(t) \leq 0$ .

Using the expression of  $N'_h(t)$  from (10) we obtain

$$\Lambda_h = f_h \left( S_e^{**} + E_e^{**} + I_e^{**} + S_a^{**} + E_a^{**} + I_a^{**} + R_a^{**} \right) + \gamma_e I_e^{**} + \gamma_a I_a^{**}.$$
(16)

When we put (5) and (16) in (10), it implies that

$$\begin{split} N'_{h}(t) &= -f_{h} \Big( V_{se}(t) + V_{ee}(t) + V_{ie}(t) + V_{sa}(t) + V_{ea}(t) + V_{ia}(t) + V_{ra}(t) \Big) \\ &- \gamma_{e} V_{ie}(t) - \gamma_{a} V_{ia}(t). \end{split}$$

It follows that

$$\begin{aligned} \mathcal{V}_{1}'(t) &= -f_{h} \Big( V_{se}(t) + V_{ee}(t) + V_{ie}(t) + V_{sa}(t) + V_{ea}(t) + V_{ia}(t) + V_{ra}(t) \Big)^{2} \\ &- \Big( V_{se}(t) + V_{ee}(t) + V_{ie}(t) + V_{sa}(t) + V_{ea}(t) + V_{ia}(t) + V_{ra}(t) \Big) \\ &\times \Big( \gamma_{e} V_{ie}(t) + \gamma_{a} V_{ia}(t) \Big). \end{aligned}$$

Therefore,  $\mathcal{V}_1'(t) \leq 0$ . Moreover,

$$\begin{aligned} E'_{m}(t) + I'_{m}(t) &= -k_{m}A^{*} - k_{m}\big(E_{m}(t) + I_{m}(t)\big) - f_{m}\big(E_{m}(t) + I_{m}(t)\big) \\ &= -k_{m}\big(V_{em}(t) + V_{im}(t)\big) + k_{m}S^{**}_{m} - f_{m}\big(E_{m}(t) + I_{m}(t)\big) \\ &\leq -k_{m}\big(V_{em}(t) + V_{im}(t)\big) + k_{m}S^{**}_{m}. \end{aligned}$$

It then follows that

$$\begin{aligned} \mathcal{V}_{2}'(t) &\leq \left( V_{em}(t) + V_{im}(t) \right) \times \left( -k_{m} \left( V_{em}(t) + V_{im}(t) \right) + k_{m} S_{m}^{**} \right) \\ &\leq -k_{m} \left( V_{em}(t) + V_{im}(t) \right)^{2} + k_{m} S_{m}^{**} \left( V_{em}(t) + V_{im}(t) \right) \\ &\leq -k_{m} \left( V_{em}(t) + V_{im}(t) \right)^{2} \\ &\leq 0, \end{aligned}$$

since  $V_{em}(t) \leq 0$  and  $V_{im}(t) \leq 0$  on  $\Delta^{**}$ .

Hence,  $\mathcal{V}'(t) \leq 0$  and then LaSalle's invariant principle [20] implies that the endemic equilibrium is globally asymptotically stable on  $\Delta^{**}$ .

*Remark* 3.2 The local stability of the equilibrium point  $DFE_0$  can be established using the mosquitoes growth rate r. We have previously obtained

$$\mathcal{R}_0 = \sqrt{K_{27}K_{72} + K_{47}K_{74} + K_{57}K_{75}},$$

thus,

$$\mathcal{R}_0 = f_\kappa(r) = \sqrt{\kappa \left(1 - \frac{1}{r}\right)},$$

where

$$\begin{split} \kappa &= \frac{\nu_e}{B^*C^*} \times c_{me}n_a \frac{S_e^*}{N_h^*} \times \frac{\nu_m}{H^*} \times \frac{1}{f_m^2} \times c_{em}n_a \frac{s_P K_P}{\chi_P N_h^*} \\ &+ \frac{\nu_a}{D^*E^*} \times c_{ma}n_a \frac{S_a^*}{N_h^*} \times \frac{\nu_m}{H^*} \times \frac{1}{f_m^2} \times c_{am}n_a \frac{s_P K_P}{\chi_P N_h^*} \\ &+ \frac{\alpha_a \nu_a}{D^*E^*G^*} \times c_{ma}n_a \frac{S_a^*}{N_h^*} \times \frac{\nu_m}{H^*} \times \frac{1}{f_m^2} \times \tilde{c}_{am}n_a \frac{s_P K_P}{\chi_P N_h^*}. \end{split}$$

*Remark* 3.3 Suppose that r > 1. Hence, the function  $f_{\kappa}$  is continuous and derivable. Moreover, some easy calculations give

$$f_{\kappa}'(r) = \frac{\kappa}{r^2 \sqrt{\kappa(1-\frac{1}{r})}}.$$

It is clear that  $f'_{\kappa}(r)$  is positive for r > 1; it follows that the larger the threshold r is, the larger the basic reproduction number  $\mathcal{R}_0$  becomes.

**Lemma 3.2** Let consider the following threshold parameter:  $r_0 = \frac{\kappa}{\kappa-1}$ ,  $\kappa \neq 1$ .

- (i)  $\mathcal{R}_0 < 1$  is equivalent to  $1 < r < r_0$ .
- (ii)  $\mathcal{R}_0 > 1$  is equivalent to  $r > r_0$ .

Proof Indeed, we have

(i) 
$$\mathcal{R}_0 < 1 \quad \Leftrightarrow \quad f_{\kappa} < 1 \quad \Leftrightarrow \quad \kappa \left(1 - \frac{1}{r}\right) < 1 \quad \Leftrightarrow \quad r < \frac{\kappa}{\kappa - 1} = r_0, \quad \kappa \neq 1,$$

and

(ii) 
$$\mathcal{R}_0 > 1 \quad \Leftrightarrow \quad f_{\kappa} > 1 \quad \Leftrightarrow \quad \kappa \left(1 - \frac{1}{r}\right) > 1 \quad \Leftrightarrow \quad r > \frac{\kappa}{\kappa - 1} = r_0, \quad \kappa \neq 1.$$

#### Theorem 3.5

- (i) If  $r \le 1$  then the disease-free equilibrium  $(0,0,0,0,S_e^*,0,0,S_a^*,0,0,0,0,0)$  of the system (9) is globally asymptotically stable.
- (ii) If  $1 < r < r_0$  then the disease-free equilibrium point ( $E^*, L^*, P^*, A^*, S_e^*, 0, 0, S_a^*, 0, 0, 0, 0, 0$ ) of the system (9) is globally asymptotically stable.

(iii) If  $r > r_0$  then the endemic equilibrium  $(E^*, L^*, P^*, A^*, S_e^{**}, E^{**}, I_e^{**}, S_a^{**}, E_a^{**}, I_a^{**}, R_a^{**}, E_m^{**}, I_m^{**})$  of the system (9) is globally asymptotically stable.

*Proof* The key idea to prove this theorem is to use the theory of internally chain transitive sets [21, 36].

Let  $\Phi(t)$  be the solution of the system (9) on  $\Omega_m \times \Omega_h$ , that is,  $\Phi(t)(E(0), L(0), P(0), S_e(0), E_e(0), I_e(0), S_a(0), E_a(0), I_a(0), R_a(0), A(0), E_m(0), I_m(0))$ . Then  $\Phi(t)$  is compact for each t > 0. Let  $\omega = \omega(E(0), L(0), P(0), A(0), S_e(0), E_e(0), I_e(0), S_a(0), E_a(0), I_a(0), R_a(0), E_m(0), I_m(0))$  be the omega limit set of  $\Phi(t)(E(0), L(0), P(0), S_e(0), E_e(0), I_e(0), S_a(0), E_a(0), I_a(0), R_a(0), A(0), R_a(0), P(0), S_e(0), E_e(0), I_e(0), S_a(0), E_a(0), I_a(0), R_a(0), A(0), E_m(0), I_m(0))$  be the omega limit set of  $\Phi(t)(E(0), L(0), P(0), S_e(0), E_e(0), I_e(0), S_a(0), E_a(0), I_a(0), R_a(0), A(0), E_m(0), I_m(0))$  be the omega limit set of  $\Phi(t)(E(0), L(0), P(0), S_e(0), E_e(0), I_e(0), S_a(0), E_a(0), I_a(0), R_a(0), A(0), E_m(0), I_m(0))$  It then follows that  $\omega$  is an internally chain transitive set for  $\Phi(t)$ .

(i) In the case where  $r \leq 1$ , then

,

$$E(t) \to 0$$
,  $L(t) \to 0$ ,  $P(t) \to 0$  and  $A(t) \to 0$  as  $t \to +\infty$ .

Since  $A(t) \to 0$  as  $t \to +\infty$ ,  $E_m(t) \to 0$  and  $I_m(t) \to 0$  as  $t \to +\infty$ . Thus, we have  $\omega = (0, 0, 0, 0, 0, 0) \times \omega_1$  with  $\omega_1 \subset \mathbb{R}^7$ .

Moreover, we have  $\Phi(t)|_{\omega}(0,0,0,0,0,0,0,S_e^*,0,0,S_a^*,0,0,0) =$ 

 $(0, 0, 0, 0, 0, 0, 0, \Phi_1(t)(S_e(0), E_e(0), I_e(0), S_a(0), E_a(0), I_a(0), R_a(0)))$  associated with the following system:

$$S'_{e}(t) = p\Lambda_{h} - f_{h}S_{e}(t),$$

$$E'_{e}(t) = -(f_{h} + v_{e})E_{e}(t),$$

$$I'_{e}(t) = v_{e}E_{e}(t) - (f_{h} + \gamma_{e} + \alpha_{e})I_{e}(t),$$

$$S'_{a}(t) = (1 - p)\Lambda_{h} + \beta_{a}R_{a}(t) - f_{h}S_{a}(t),$$

$$E'_{a}(t) = -(f_{h} + v_{a})E_{a}(t),$$

$$I'_{a}(t) = v_{a}E_{a}(t) - (f_{h} + \gamma_{a} + \alpha_{a})I_{a}(t),$$

$$R'_{a}(t) = \alpha_{e}I_{e}(t) + \alpha_{a}I_{a}(t) - (f_{h} + \beta_{a})R_{a}(t).$$
(17)

From the second and the fifth equation of the system (17), we have  $E_e(t) \rightarrow 0$  and  $E_a(t) \rightarrow 0$  as  $t \rightarrow +\infty$ . Using the limit system of the system (17), it then follows that  $I_e(t) \rightarrow 0$  and  $I_a(t) \rightarrow 0$ . Hence, we deduce from the last equation that  $R_a(t) \rightarrow 0$ . Finally, we obtain the following limit system:

$$S'_{e}(t) = p\Lambda_{h} - f_{h}S_{e}(t),$$

$$S'_{a}(t) = (1-p)\Lambda_{h} - f_{h}S_{a}(t).$$
(18)

It is easy to see from the above system that  $S_e(t) \rightarrow 0$  and  $S_a(t) \rightarrow 0$  as  $t \rightarrow 0$ . (ii) In the case where r > 1, then from Theorem 2.5, we have

$$E(t) \to E^*$$
,  $L(t) \to L^*$ ,  $P(t) \to P^*$ , and  $A(t) \to A^*$  as  $t \to +\infty$ ,

for any E(0) > 0, L(0) > 0, P(0) > 0, A(0) > 0. Hence, we have  $\omega = (E^*, L^*, P^*, A^*) \times \omega_2$  with  $\omega_2 \subset \mathbb{R}^9$ . It is easy to see that  $\Phi(t)|_{\omega}(E^*, L^*, P^*, A^*, S_e^*, 0, 0, S_a^*, 0, 0, 0, 0, 0) =$  $(E^*, L^*, P^*, A^*, \Phi_2(t)(S_e(0), E_e(0), I_e(0), S_a(0), E_a(0), I_a(0), R_a(0), E_m(0), I_m(0)))$ associated with the system (12).

Since  $\omega$  is an internally limit set for  $\Phi(t)$ , it is easy to see that  $\omega_2$  is an internally chain transitive set for  $\Phi_2(t)$ . Since  $r < r_0$ , thanks to Theorem 3.2, the disease-free equilibrium  $(S_e^*, 0, 0, S_a^*, 0, 0, 0, 0, 0)$  is globally asymptotically stable for the limit system (12). It then follows from Theorem 3.2 and Remark 4.6 in [16] that  $\omega_2 = \{(S_e^*, 0, 0, S_a^*, 0, 0, 0, 0, 0)\}$  and  $\omega = \{(E^*, L^*, P^*, A^*, S_e^*, 0, 0, S_a^*, 0, 0, 0, 0, 0, 0)\}$ . Hence, if  $1 < r < r_0$ , then the disease-free equilibrium  $(E^*, L^*, P^*, A^*, S_e^*, 0, 0, S_a^*, 0, 0, 0, 0, 0, 0)$  is globally asymptotically stable through the system (9).

(iii) In the case where  $r > r_0$ , thanks to Theorem 2.5, we have

 $E(t) \to E^*, \qquad L(t) \to L^*, \qquad P(t) \to P^* \text{ and } A(t) \to A^* \text{ as } t \to +\infty,$ 

for any E(0) > 0, L(0) > 0, P(0) > 0, A(0) > 0.

Hence, we have  $\omega = \{(E^*, L^*, P^*, A^*)\} \times \omega_3$  with  $\omega_3 \subset \mathbb{R}^9$  and  $\Phi(t)|_{\omega}(E^*, \dots, A^*, S_e(0), \dots, I_m(0)) = (E^*, \dots, A^*, \Phi_2(t)(S_e(0), \dots, I_m(0)))$  where  $\Phi_2(t)$ 

is the solution semiflow of the system (12). Thanks to Lemma 3.2,  $r > r_0$ , implies that  $\mathcal{R}_0 > 1$  and then  $\omega_2 \neq 0_{\mathbb{R}^9}$ .

Since  $(S_e^{**}, E_e^{**}, I_e^{**}, S_a^{**}, E_a^{**}, I_a^{**}, R_a^{**}, E_m^{**}, I_m^{**})$  is globally asymptotically stable for the system (17) in  $\mathbb{R}^{*9}$ ,  $\omega_3 \cap W^s((S_e^{**}, E_e^{**}, I_e^{**}, S_a^{**}, E_a^{**}, I_a^{**}, R_a^{**}, E_m^{**}, I_m^{**})) \neq \emptyset$ . Hence, the statement (iii) is valid.

## **4** Numerical simulations

In this section we perform some numerical results in order to illustrate theoretical results which were previously established. Our numerical simulation will be performed using the MATLAB technical computing software with the fourth-order Runge–Kutta method [1, 23, 26, 30]. The values of the parameters are given in Table 7.

#### 4.1 Dynamical model for vector population growth

Firstly, using the following initial conditions: E(0) = 35, L(0) = 25, P(0) = 30, A(0) = 45 and the mosquito's parameter values for extinction given in Table 7, we obtain Fig. 3. These values lead to the condition  $r \le 1$ ; that is, the mosquitoes' population disappears. This result confirms Theorem 2.2.

Secondly, using the following initial conditions: E(0) = 50, L(0) = 40, P(0) = 8, A(0) = 6 and the numerical values of parameters in Table 7 which lead to the threshold *r* greater than unity, we get Fig. 4. We observe that when *r* is greater than unity, the mosquitoes' population persists. This result supports Theorem 2.3.

These two observations show that the threshold parameter r may provide conditions in order to control the proliferation of the mosquito population.

#### 4.2 Global model of malaria transmission

The initial conditions used here are E(0) = 50, L(0) = 40, P(0) = 30, A(0) = 80,  $S_e(0) = 50$ ,  $S_a(0) = 200$ ,  $I_e(0) = 500$ ,  $S_a(0) = 25$ ,  $E_a(0) = 150$ ,  $I_a(0) = 350$ ,  $R_a(0) = 400$ ,  $S_m(0) = 50$ ,  $E_m(0) = 15$  and  $I_m(0) = 100$ . Using the above initial conditions and the numerical values of human parameters in Table 7, which lead to  $\mathcal{R}_0$  less than unity, we get Fig. 5. This

Parameters	Value for extinction	Value for persistence	Reference	Dimension
p	0.25	0.8	estimated	dimensionless
C <sub>me</sub>	0.021	0.03	[13]	dimensionless
Cem	0.11	0.45	[13]	dimensionless
C <sub>ma</sub>	0.012	0.022	[13]	dimensionless
Cam	0.08	0.35	[13]	dimensionless
<i>Č</i> am	0.008	0.002	[13]	dimensionless
$\nu_e$	0.10	0.10	[13]	/days
$v_a$	0.06	0.09	[13]	/days
$v_m$	0.083	0.083	[13]	/days
$\alpha_e$	0.001	0.001	[13]	/days
$\alpha_a$	0.01	0.01	[13]	/days
$\gamma_e$	0.000018	0.000018	[13]	/days
$\gamma_a$	0.00003	0.00003	[13]	/days
$\beta_a$	0.0055	0.0027	[13]	/days
$\Lambda_h$	50	85	estimated	humans/week
f <sub>h</sub>	0.00063	0.00063	[13]	/human/days
f <sub>m</sub>	0.1	0.1	[13]	/mosquito/days
na	0.25	0.5	[13]	human/days
K <sub>E</sub>	10,000	10,000	estimated	space
KL	5000	5000	estimated	space
K <sub>P</sub>	4000	4000	estimated	space
Ь	2	10.7	[1]	/days
SE	0.6	0.4	[1]	dimensionless
d <sub>E</sub>	0.3	0.36	[1]	dimensionless
SL	0.4	0.5	[1]	dimensionless
$d_L$	0.3	0.34	[1]	dimensionless
Sp	0.25	0.3	[1]	dimensionless
d <sub>P</sub>	0.15	0.17	[1]	dimensionless
f <sub>m</sub>	0.6	0.15	[1]	dimensionless

 Table 7
 Human and vector parameter's values for the malaria model



figure shows the extinction of the different infected classes of global disease dynamics model. These observations confirm Theorem 3.1. With the same above initial conditions for vectors, taking  $S_e(0) = 50$ ,  $S_a(0) = 200$ ,  $I_e(0) = 500$ ,  $S_a(0) = 25$ ,  $E_a(0) = 150$ ,  $I_a(0) = 350$ ,





 $R_a(0) = 400, S_m(0) = 50, E_m(0) = 15, I_m(0) = 100$  for humans and the above numerical values of human parameters in Table 7, which lead to  $\mathcal{R}_0$  greater than unity, we get Fig. 6. This figure shows the persistence of the different infected classes of global disease dynamics model. These observations confirm Theorem 3.2.

### **5** Discussions and conclusion

In this paper, we have proposed mathematical models to describe the vector population growth global dynamics and the malaria virus transmission to human population.



In the first part, we have proposed the vector population dynamics model including auto-regulation phenomena of eggs, larvae and pupae. We used the Verhulst–Pearl logistic functions in order to gain insight into its qualitative features. For this model, we found that the mosquito growth rate r is the threshold condition for the existence of the endemic state. Besides, for r greater than unity, we proved using the Lyapunov function that the endemic equilibrium is globally asymptotically stable. The study of this model shows that the effect of immature stages is very important on the mosquitoes' population proliferation [1, 23, 30].

Moreover, we have proposed a model to describe the malaria virus transmission to human population including some biological complexities as human host susceptibility. We divide the human hosts into two major types: the first type is called non-immune and comprised all humans who have not acquired the immunity against malaria; the second type is called semi-immune and represents all the people who have at least once acquired immunity during his life. For a better understanding of the mosquitoes population proliferation effect on the malaria disease spread, the two models were associated and the global study done. For this global model, the common basic reproduction number was determined using the next generation matrix idea [17, 18, 32]. As a further new insight, there was established a very interesting relationship between the common basic reproduction number and the regulatory threshold parameters of mosquito population, r, and its implications for malaria management analyzed. We found another threshold parameter, called  $r_0$ , which is expressed using the two model parameters. Using the threshold  $r_0$ , we established the global transmission model stability.

Finally, numerical simulations are carried out to support all the above theoretical results and provide conditions in order to control the proliferation of mosquito population and its implication on malaria spread control. It shows clearly that malaria management is concerned firstly lowering the mosquito threshold parameters to a value less than unity. This condition leads to the mosquito population's disappearance. This is an ideal case in the fight against malaria but it can have some environment mistakes. In the other case, keeping the threshold r between 1 and  $r_0$  leads to the basic reproduction number less than unity, so the disease disappears.

In this study, we consider a homogeneously mixed population; however, each individual may have a heterogeneous number of contacts in the population. Meanwhile, the contact network structure is ignored. For future work, it would be fair to include the network structure as in [33, 34], which will make the model more realistic.

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#### Competing interests

The authors declare that they have no competing interests.

#### Authors' contributions

All authors worked together to produce the results and read and approved the final manuscript.

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