

Mathematical model of the aquatic phase in the population growth dynamics of *Aedes aegypti* contaminated with *Wolbachia*

Modelo matemático de la fase acuática en la dinámica de crecimiento de la población de *Aedes aegypti* contaminada con *Wolbachia*

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ABSTRACT

Keywords:

Aedes aegypti, water phase, mathematical modeling, *Wolbachia*, ordinary differential equations.

Dengue is an infectious disease of global public health significance and is a leading cause of death. A study focusing on the aquatic phase (egg, larva and pupa) of the *Aedes aegypti* mosquito, with and without *Wolbachia*, is being conducted. A mathematical model is formulated, the results suggest that the intrinsic ovoposition rate is the parameter that most affects the non-trivial equilibrium solution. As a conclusion, it is suggested that, under certain conditions, the reproduction of *Wolbachia*-contaminated mosquitoes in the aquatic phase is small compared to the reproduction of wild mosquitoes, which could affect the success of the control strategy.

RESUMEN

Palabras clave:

Aedes aegypti, fase acuática, modelización matemática, *Wolbachia*, ecuaciones diferenciales ordinarias.

El dengue es una enfermedad infecciosa de importancia para la salud pública mundial y es una de las principales causas de muerte. Se realiza un estudio centrado en la fase acuática (huevo, larva y pupa) del mosquito *Aedes aegypti*, con y sin *Wolbachia*. Se formula un modelo matemático, cuyos resultados sugieren que la tasa de ovoposición intrínseca es el parámetro que más afecta a la solución de equilibrio no trivial. Como conclusión, se sugiere que, bajo ciertas condiciones, la reproducción de los mosquitos contaminados con *Wolbachia* en la fase acuática es pequeña en comparación con la reproducción de los mosquitos salvajes, lo que podría afectar al éxito de la estrategia de control.

Introduction

There is an extensive literature on mathematical modeling of the population dynamics of the *Aedes aegypti* mosquito that considers both the aquatic and aerial phases of the mosquito [1-9]. However, most references to aquatic phase modeling consider a population group that compacts eggs, larvae and pupae into a single dynamic variable. This form of modeling does not consider factors such as the percentage of eggs that develop into larvae or the percentage of larvae that develop into pupae. Several studies have shown that the aquatic phase is fundamental,

not only in the growth process of the vector's population, but also in the epidemiology of mosquito-borne diseases. Moreover, around the world, the main strategies for controlling the mosquito population, such as mosquito breeding ground elimination campaigns, focus on reducing the population of the mosquito in its aquatic phase. However, there are other control strategies such as biological ones that contribute significantly to the reduction of the basic breeding number of the vector. All this motivates us to formulate a mathematical model that describes the dynamics of the aquatic phase explicitly considering three population groups in the aquatic phase.

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Methodology

Mathematical model –

The dynamics consider natural (wild) mosquito populations and mosquitos contaminated with *Wolbachia*. For each group, the life of mosquitoes is defined and divided into the following stages: eggs, larvae and pupae. The following variables are defined on the basis of the above:

Aquatic phase for wild or natural mosquitoes:

$E(t)$ is the amount of eggs in time t .

$L(t)$ is the amount of larvae in time t .

$P(t)$ is the amount of pupae in time t .

Aquatic phase for mosquitoes contaminated with *Wolbachia*:

a) $E_w(t)$ is the amount of eggs with *Wolbachia* in time t .

b) $L_w(t)$ is the amount of larvae with *Wolbachia* in time t .

c) $P_w(t)$ is the amount of pupae with *Wolbachia* over time t .

The growth dynamics for the wild mosquito egg population is modeled by the following differential equation (1):

$$\frac{dE}{dt} = \phi + \phi_w(1 - \xi) - (\gamma_1 + \mu_E)E \quad (1)$$

Where (ϕ) is the intrinsic ovoposition rate, $(\phi_w = \phi a)$ is the ovoposition rate of mosquitoes with *Wolbachia*, with $0 < a < 1$ a rate of adjustment that arises because not all the eggs of adult females with *Wolbachia* hatch, (ξ) is the probability of the *Wolbachia* bacteria being transferred vertically, (γ_1) is the per capita rate of transformation of the egg to the larval stage, (μ_E) is the natural death rate of eggs.

The dynamics for eggs with *Wolbachia* is described by the following differential equation (2):

$$\frac{dE_w}{dt} = \phi_w \xi - (\gamma_1 + \mu_{E_w})E_w \quad (2)$$

Where (μ_{E_w}) is the natural death rate of eggs with

Wolbachia.

The dynamics for wild mosquito larvae is modeled by Equation (3):

$$\frac{dL}{dt} = \gamma_1 E - \mu_L L - \theta L \quad (3)$$

Where (θ) is the per capita rate of transformation of the larva to the pupal stage.

The dynamics for *Wolbachia*-contaminated mosquito larvae is modeled by Equation (4):

$$\frac{dL_w}{dt} = \gamma_1 E_w - \mu_{L_w} L_w - \theta L_w \quad (4)$$

Where (μ_{L_w}) is the natural death rate of the larvae with *Wolbachia*.

The dynamics for wild mosquito pupae is modeled by Equation (5):

$$\frac{dP}{dt} = \theta L - \mu_P P \quad (5)$$

Where (μ_{P_w}) is the natural death rate of *Wolbachia* pupae.

Be $x(t) = (E(t), E_w(t), L(t), L_w(t), P(t), P_w(t))^T$ a solution of the system of equations (1) to (6), then $x \in \Omega$, where:

$$\Omega = \{x \in \mathbb{R}^6 : N \leq (\phi + \phi_w) / \mu\} \quad (6)$$

Where $N = E + E_w + L + L_w + P + P_w$ and $\mu = \min \{\mu_E, \mu_{E_w}, \mu_L, \mu_{L_w}, \mu_P, \mu_{P_w}\}$, the whole Ω defined in (7) is the set of biological interest since any solution of the system (1)-(6) which starts in Ω stays there for everything $t \geq 0$. The system (1)-(6) has a single globally asymptotically stable equilibrium solution given by Equation (8):

$$x^* = (E^*, E_w^*, L^*, L_w^*, P^*, P_w^*)^T \quad (7)$$

Where,

$$E^* = \frac{\phi[1+a(1-\xi)]}{\gamma_1+\mu_E}, E_w^* = \frac{a\phi\xi}{\gamma_1+\mu_{Ew}}, L^* = \frac{\gamma_1}{\theta+\mu_L} \frac{\phi[1+a(1-\xi)]}{\gamma_1+\mu_E},$$

$$L_w^* = \frac{\gamma_1}{\theta+\mu_{Lw}} \frac{a\phi\xi}{\gamma_1+\mu_{Ew}}, P^* = \frac{\theta}{\mu_P} \frac{\gamma_1}{\theta+\mu_L} \frac{\phi[1+a(1-\xi)]}{\gamma_1+\mu_E}, P_w^* = \frac{\theta}{\mu_{Pw}} \frac{\gamma_1}{\theta+\mu_{Lw}} \frac{a\phi\xi}{\gamma_1+\mu_{Ew}}.$$

The above implies that any solution of the system (1)-(6) in Ω tends to approach x^* equilibrium as time t grows.

Result and Discussion

Numerical simulations

A parameter sensitivity analysis similar to that performed by other authors [8-10] verifies the intrinsic ovoposition rate, (ϕ) , is the parameter that most influences the magnitude of the equilibrium solution (x^*). According to Equation (8) it can be seen that all the populations are directly proportional (ϕ) , which implies that $(x^*=\phi X)$ where (X) is a vector that depends on the other parameters. On the other hand, after (ϕ) , the parameters that most affect the sensitivity of (x^*) are the rate of adjustment (a) , and the probability of vertical transmission of bacteria (ξ) . Following the order is the per capita rate of transformation of the egg to the larval stage (γ_1) and the per capita rate of transformation of the larva to the pupal stage (θ) . Finally, at the last level are the mortality rates. There are many climatic variables (temperature, rainfall and humidity, among others) focuses on temperature or precipitation. So, Table 1, presents a range of estimated parameters based on temperature changes [6].

Table 1. Description of parameters

| Parameter | Range | Reference value | Reference |
|------------|--------------|-----------------|-----------|
| ϕ | (0.00-1500) | 1.840 | [6] |
| a | (0.00 -1.00) | 0.800 | Estimated |
| ϕ_w | (0.00-1500) | 1.120 | Estimated |
| ξ | (0.00 -0.30) | 0.007 | [6] |
| θ | (0.08 -0.35) | 0.140/day | [6] |
| γ_1 | (0.10 -0.50) | 0.400/day | [6] |
| μ_E | (0.07 -0.30) | 0.200/day | [6] |
| μ_{Ew} | (0.00 -1.00) | 0.200/day | Estimated |
| μ_L | (0.07 -0.30) | 0.180/day | [6] |
| μ_{Lw} | (0.00 -1.00) | 0.180/day | Estimated |
| μ_P | (0.07 -0.25) | 0.170/day | [6] |
| μ_{Pw} | (0.00 -1.00) | 0.170/day | [6] |

The parameter ranges $(\phi, \xi, \theta, \gamma_1, \mu_E, \mu_L, \mu_P)$ were estimated using functions adjusted to data from Chiang Mia province in Thailand [6]. Since no references were found to the values of $(a, \phi_w, \mu_{Ew}, \mu_{Lw}$ and $\mu_{Pw})$ were considered the broadest ranges. So, Figure 1, shows the plots of eggs, larvae and pupae, uncontaminated and contaminated with *Wolbachia* according to the ovoposition rate (ϕ) , as it is observed for each egg contaminated with *Wolbachia* there are 300 eggs without *Wolbachia*, the same relation is presented with the larvae and pupae. Figure 2, shows the plots of eggs, larvae and pupae, uncontaminated and contaminated with *Wolbachia* as a function of the rate of adjustment (a) , it is observed that under the set of reference values (Table 1) the variation of the population of eggs, larvae and

pupae with *Wolbachia* is very small in relation to the population of eggs, larvae and pupae without *Wolbachia*. Figure 3, shows the graphs of eggs, larvae and pupae, uncontaminated and contaminated with *Wolbachia* as a function of the ovoposition rate (ϕ) and the adjustment rate (a), it is observed that the yield in all populations increased in relation to the yields with respect to each parameter.

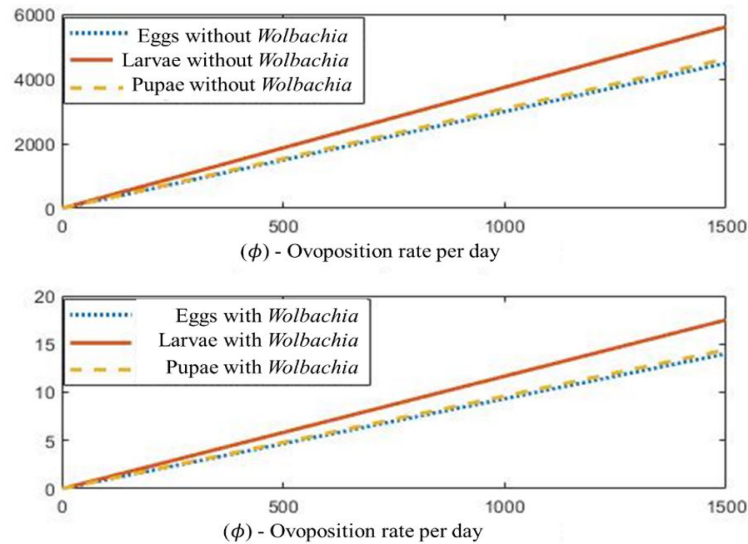


Figure 1. a) Population growth of eggs, larvae and pupae, without *Wolbachia*, with respect to the ovoposition rate (ϕ), taking the range of (ϕ) and the fixed reference values for the remaining parameters. b) Population growth of eggs, larvae and pupae, with *Wolbachia*, with respect to the ovoposition rate (ϕ), taking the range of (ϕ) and the fixed reference values for the remaining parameters.

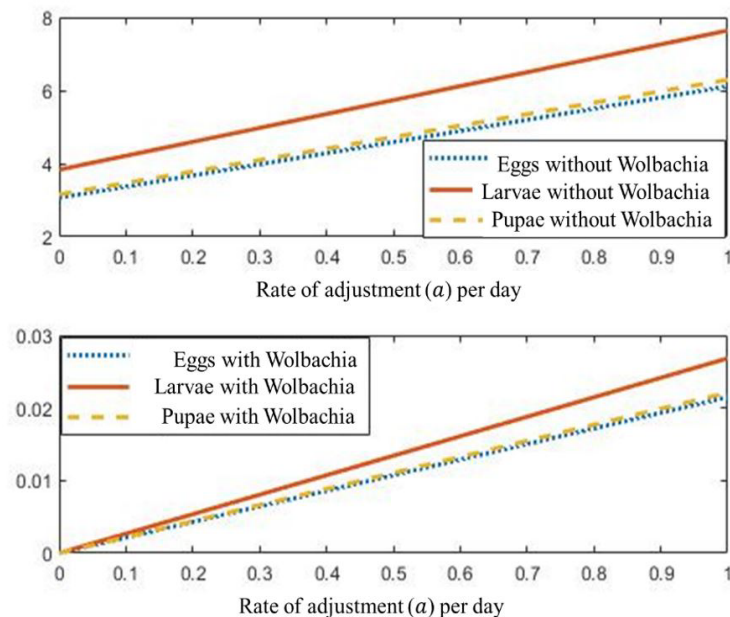


Figure 2. a) Growth of the population of eggs, larvae and pupae, without *Wolbachia*, respect to the adjustment rate (a), taking the range of the adjustment rate (a) and the reference values, fixed for the rest of parameters. b) Growth of the population of eggs, larvae and pupae, with *Wolbachia*, respect to the adjustment rate (a), taking the range of the

adjustment rate (a) and the reference values, fixed for the rest of parameters.

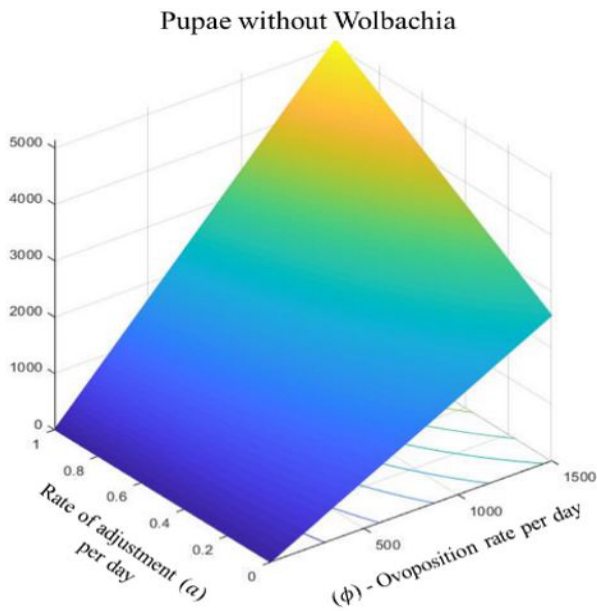


Figure 3. a) Growth of the population of pupae without *Wolbachia*, respect to the adjustment rate (a) and the ovoposition rate (ϕ).

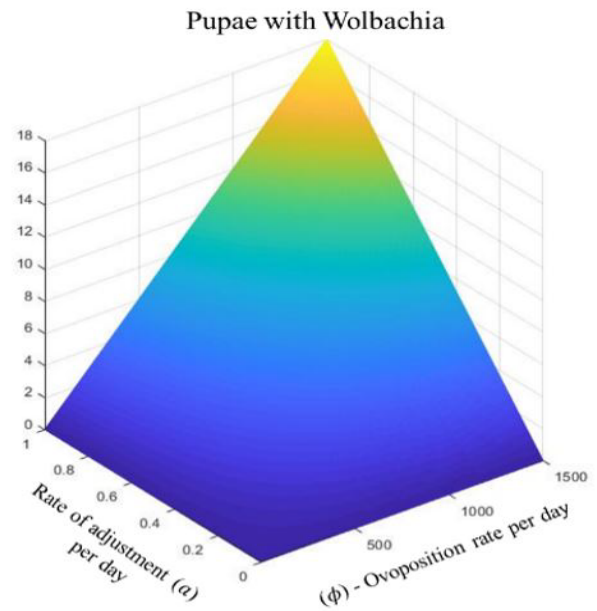


Figure 3. b) Growth of the population of pupae with *Wolbachia*, respect to the adjustment rate (a) and the ovoposition rate (ϕ).

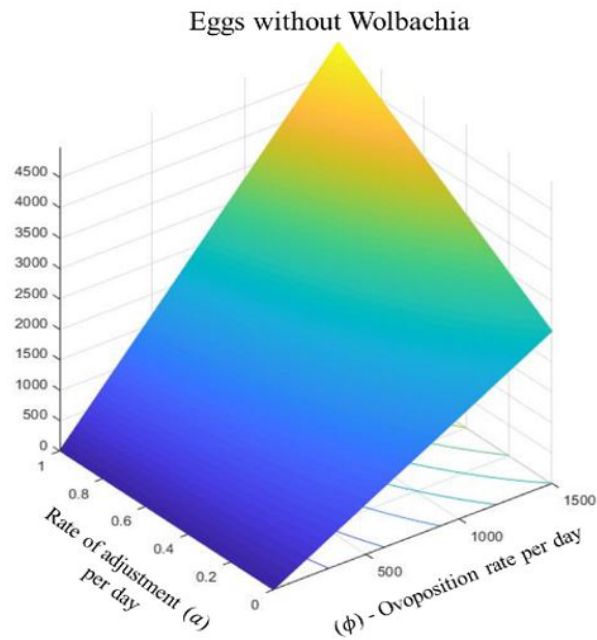


Figure 3. c) Growth of the population of eggs without *Wolbachia*, respect to the adjustment rate (a) and the ovoposition rate (ϕ).

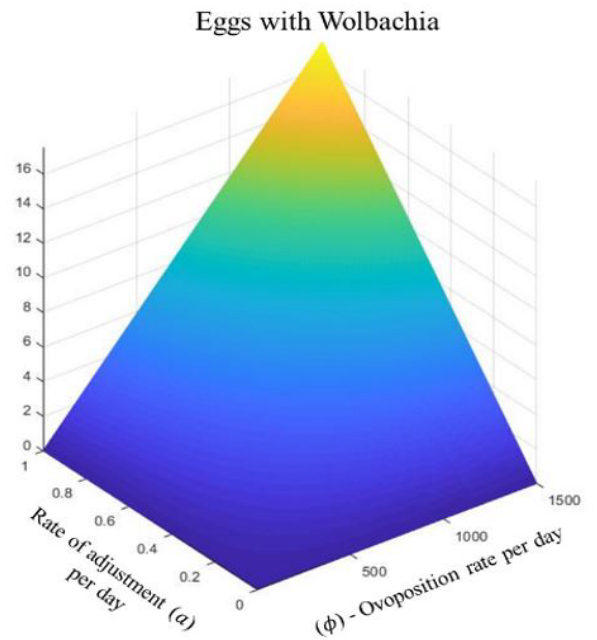


Figure 3. d) Growth of the population of eggs with *Wolbachia*, respect to the adjustment rate (a) and the ovoposition rate (ϕ).

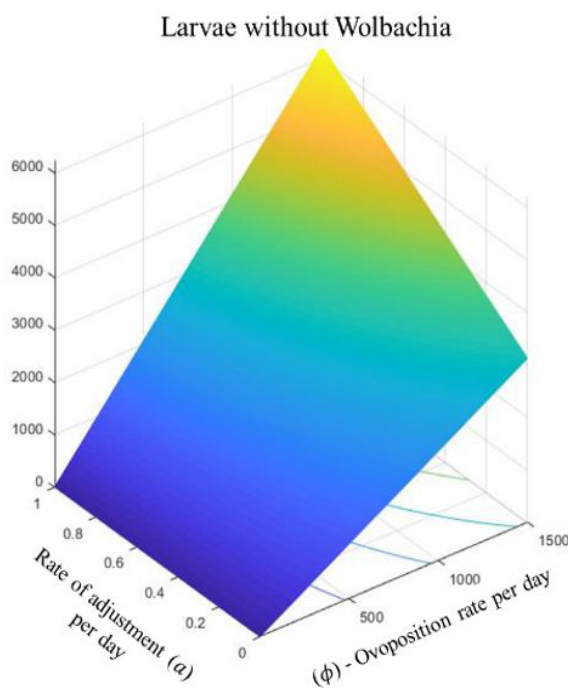


Figure 3. e) Growth of the population of larvae without *Wolbachia*, respect to the adjustment rate (a) and the ovoposition rate (ϕ).

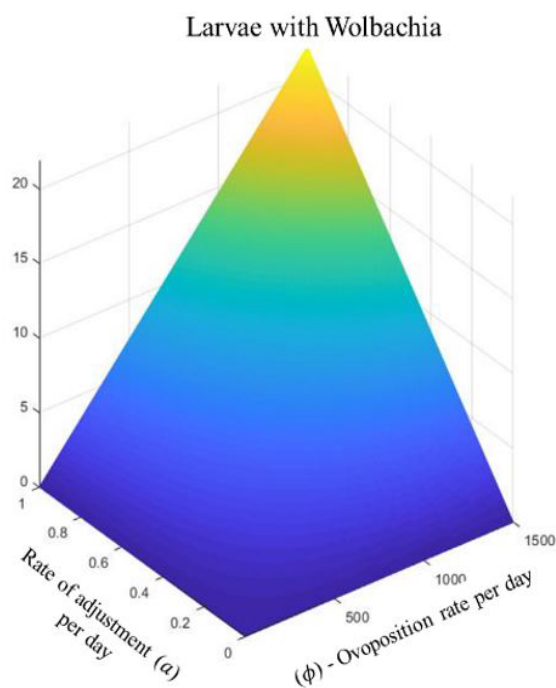


Figure 3. f) Growth of the population of larvae with *Wolbachia*, respect to the adjustment rate (a) and the ovoposition rate (ϕ).

Figure 3. Growth of the population of eggs, larvae and pupae, with and without *Wolbachia*, with respect to the adjustment rate (a) and the ovoposition rate (ϕ).

Conclusion

Within the dynamic spread of the *Aedes aegypti* mosquito and transmission of Dengue, the water phase is of great importance. In fact, the most widely used strategies to control the spread of the mosquito are aimed at reducing the population of eggs, larvae and pupae (larvicides, insecticides, elimination of breeding sites). In recent years, strategies based on biological control have been implemented, in particular the replacement of wild mosquitoes by *Wolbachia*-infected mosquitoes is a strategy that is gaining ground among all biological control strategies. Although the method concentrates on the aerial phase of the mosquito, the aquatic phase involves the breeding of *Wolbachia*-contaminated eggs, larvae and pupae, which will later develop into contaminated adult mosquitoes. In this sense, the amount of contaminated mosquitoes that pass from the aquatic phase to the aerial phase contributes directly to the success of the strategy. The results of this work suggest that under certain conditions the breeding of *Wolbachia*-contaminated mosquitoes in the aquatic phase is very small compared to the breeding of wild mosquitoes. This may affect the success of the control strategy.

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