

Local Stability Analysis of Age Structural Model for Herpes Zoster in Thailand

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Abstract—Herpes zoster is a disease that manifests as a dermatological condition. The characteristic of this disease is an irritating skin rash with blisters. This is often limited to one side of body. From the data of Herpes zoster cases in Thailand, we found that age structure effects to the transmission of this disease. In this study, we construct the age structural model of Herpes zoster in Thailand. The local stability analysis of this model is given. The numerical solutions are shown to confirm the analytical results.

Keywords—Age structural model, Herpes zoster, local stability, Numerical solution.

I. INTRODUCTION

HERPES zoster is a viral disease characterized by a painful skin rash with blisters in a limited area on one side of the body (left or right), often in a stripe. The initial infection with varicella zoster virus (VZV) causes the acute, short-lived illness chickenpox which generally occurs in children and young adults. Herpes zoster is not the same disease as herpes simplex, despite the name similarity; both the varicella zoster virus and herpes simplex virus belong to the same viral subfamily *Alpha herpesviridae* [1]. This disease is also known as shingles or zoster, is a reactivated VZV infection of the sensory nerve ganglion and the peripheral nerve and its branches. Inflammation of the nerve axons results in a painful, burning sensation on the affected dermatome(s) being supplied by the peripheral nerve. In each year, there are about 1 million herpes zoster cases occur in the U.S. Herpes zoster more commonly occurs in white patients (35% higher incidence than in black patients), elderly patients (3 to 7 times higher incidence than in the general population). Some studies report a higher case in women (3.8 cases per 1,000 person-years and 2.6 cases per 1,000 person-years among men) [2]. Adults aged 60 years and older are risk of infection with this disease. Recurrent of infection with this disease increases with advancing age. The highest case occurs between 50 and 80 years old. Each person has 15-20% risk of herpes zoster infection.

The symptoms are usually followed by sensations of burning pain, itching, hyperesthesia or paresthesia. The pain may be mild to extreme in the affected dermatome, with sensations that are often described as stinging, tingling, aching, numbing or throbbing, and can be interspersed with quick

stabs of agonizing pain. In children, Herpes zoster is often painless, but older people are more severe than children. After 1-2 days or sometimes 3 weeks, the initial phase of patients is followed by the appearance of the skin rash. The pain and rash most commonly occurs on the torso. In Thailand, Appearance of Herpes zoster is increasing with the age. The chance of general people may be infected with Herpes zoster is 1.2-4.8/1,000 populations per year. The person who is older than 60 years old has the chance of infected about 7.2-11.8/1,000 populations per year. The trend of Thai Herpes zoster cases is increasing every year. The patient who is greater than 65 years old has the highest incidence rate [3]-[5]. Age is influence to the transmission of Herpes zoster. Older people who greater than 60 years has higher risk of Herpes zoster than children 5-10 times.

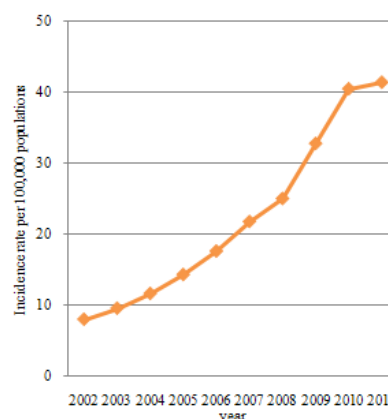


Fig. 1 Incidence rate per 100,000 populations in Thailand, 2002-2011

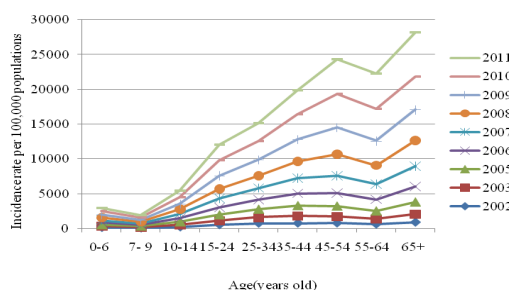


Fig. 2 Incidence rate per 100,000 populations in Thailand by age group, 2002-2011

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The data of Herpes zoster in Thailand are shown in Figs. 1 and 2. We will see that Herpes zoster cases in Thailand are

increasing every year. The highest incidence rate is found in the people greater than 65 years old.

In 1998, Allen and Thrasher [6] analyzed the model for Varicella and Herpes Zoster in United States considering vaccination in each age group and finding the appropriated parameters. In 2003, Schuette [7] studied the transmission of Varicella-zoster virus by using mathematical model. We found that basic reproductive number of Varicella-Zoster virus is calculated from the sum of basic reproduction of Varicella virus and Zoster virus. In the same year, Owoeye and Popoola [8] studied the relation between the Herpes zoster and HIV in Nigerian. They found that 50% of Herpes zoster cases are infected with HIV. In 2008, Zaman and et al. [9] studied SIR model by finding the equilibrium points and their stability conditions. The numerical solutions are evaluated by using Runge–Kutta method. Their results can be applied to many diseases such as varicella, mumps, etc. In 2010, Weinberg and et al. [10] studied HIV patients and normal people with Herpes zoster vaccination to study Herpes zoster infection. From the data of Herpes zoster in Thailand, we found that age structure effects to the transmission of this disease. In this study, we formulate the mathematical model of Herpes zoster in Thailand by considering age group of human.

II. TRANSMISSION MODEL

The transmission model of Herpes Zoster by age structure is considered. We are interested in the dynamical changes of human population with the transmission of Herpes Zoster. We separate the human population into two groups. First age group represents the human less than 65 years old. Second age group represents the human greater than 65 years old. Each human age group is divided into 6 subgroups; susceptible, exposed, infected, recovered, weak immunity and re-infected groups.

We define the variables and parameters in our model as follows:

$S_a(t)$ is the number of susceptible human of the first age group at time t ,

$E_a(t)$ is the number of exposed human of the first age group at time t ,

$I_a(t)$ is the number of infected human of the first age group at time t ,

$R_a(t)$ is the number of recovered human of the first age group at time t ,

$W_a(t)$ is the number of weak immunity human of the first age group at time t ,

$Z_a(t)$ is the number of re-infected human of the first age group at time t ,

$S_b(t)$ is the number of susceptible human of the second age group at time t ,

$E_b(t)$ is the number of exposed human of the second group at time t ,

$I_b(t)$ is the number of infected human of the second age group at time t ,

$R_b(t)$ is the number of recovered human of the second age group at time t ,

$W_b(t)$ is the number of weak immunity human of the second age group at time t ,

$Z_b(t)$ is the number of re-infected human of the second age group at time t ,

N is the total human,

N_a is the total juvenile human,

N_b is the total adult human,

a is the birth rate of human,

c_1 is the rate at which susceptible human change to be exposed human in the juvenile group,

c_2 is the rate at which exposed human change to be infectious human in the juvenile group,

c_3 is the rate at which infectious human change to be recovered human in the juvenile group,

c_4 is the rate at which recovered human change to be weak immunity human in the juvenile group,

c_5 is the rate at which weak immunity human change to be re-infected human in the juvenile group,

c_6 is the rate at which weak immunity human change to be recovered human in the juvenile group,

e_1 is the rate at which susceptible human change to be exposed human in the adult group,

e_2 is the rate at which exposed human change to be infectious human in the adult group,

e_3 is the rate at which infectious human change to be recovered human in the adult group,

e_4 is the rate at which recovered human change to be weak immunity human in the adult group,

e_5 is the rate at which weak immunity human change to be re-infected human in the adult group,

e_6 is the rate at which weak immunity human change to be recovered human in the adult group,

d is the death rate,

μ_h is the birth rate,

a is the proportion of juvenile human.

The dynamical equations for juvenile human are given by

$$\frac{d}{dt}S_a = a\mu_h N - (c_1 + d)S_a \quad (1)$$

$$\frac{d}{dt}E_a = c_1S_a - (c_2 + d)E_a \quad (2)$$

$$\frac{d}{dt}I_a = c_2E_a - (c_3 + d)I_a \quad (3)$$

$$\frac{d}{dt}R_a = c_3I_a + c_7Z_a + c_6W_a - (c_4 + d)R_a \quad (4) \quad \frac{d}{dt}s_a = m_h - (c_1 + d)s_a \quad (15)$$

$$\frac{d}{dt}W_a = c_4R_a - (c_5 + c_6 + d)W_a \quad (5) \quad \frac{d}{dt}e_a = c_1s_a - (c_2 + d)e_a \quad (16)$$

$$\frac{d}{dt}Z_a = c_5W_a - (c_7 + d)Z_a \quad (6) \quad \frac{d}{dt}i_a = c_2e_a - (c_3 + d)i_a \quad (17)$$

$$\text{where } N_a = S_a + E_a + I_a + R_a + W_a + Z_a \quad (7) \quad \frac{d}{dt}r_a = c_3i_a + c_7(1 - s_a - e_a - i_a - r_a - w_a) + c_6w_a - (c_4 + d)r_a \quad (18)$$

The dynamical equations for adult human are given by

$$\frac{d}{dt}S_b = (1 - a)\mu_h N - (e_1 + d)S_b \quad (8) \quad \frac{d}{dt}w_a = c_4R_a - (c_5 + c_6 + d)w_a \quad (19)$$

$$\frac{d}{dt}E_b = e_1S_b - (e_2 + d)E_b \quad (9) \quad \frac{d}{dt}s_b = \frac{(1 - a)m_h}{a} - (e_1 + d)s_b \quad (20)$$

$$\frac{d}{dt}I_b = e_2E_b - (e_3 + d)I_b \quad (10) \quad \frac{d}{dt}e_b = e_1s_b - (e_2 + d)e_b \quad (21)$$

$$\frac{d}{dt}R_b = e_3I_b + e_7Z_b + e_6W_b - (e_4 + d)R_b \quad (11) \quad \frac{d}{dt}i_b = e_2e_b - (e_3 + d)i_b \quad (22)$$

$$\frac{d}{dt}W_b = e_4R_b - (e_5 + e_6 + d)W_b \quad (12) \quad \frac{d}{dt}r_b = e_3i_b + e_7(1 - s_a - e_a - i_a - r_a - w_a) + e_6w_b - (e_4 + d)r_b \quad (23)$$

$$\frac{d}{dt}Z_b = c_5W_b - (e_7 + d)Z_b \quad (13) \quad \frac{d}{dt}w_b = e_4R_b - (e_5 + e_6 + d)w_b \quad (24)$$

$$\text{where } N_b = S_b + E_b + I_b + R_b + W_b + Z_b \quad (14) \quad \text{where } N = N_a + N_b, s_a + e_a + i_a + r_b + w_b + z_b = 1 \quad \text{and} \\ s_b + e_b + i_b + r_b + w_b + z_b = 1.$$

We suppose that the total human population and the human population for each group have constant sizes. Thus the rate of change for each human group equals to zero; i.e.

$$\frac{d}{dt}N = 0, \frac{d}{dt}N_a = 0, \frac{d}{dt}N_b = 0.$$

Then we obtain $d = \mu_h, N_a = aN, N_b = (1 - a)N$.

We normalize our dynamical equations by letting

$$s_a = S_a / N_a, e_a = E_a / N_a, i_a = I_a / N_a,$$

$$r_a = R_a / N_a, w_a = W_a / N_a, z_a = Z_a / N_a,$$

$$s_b = S_b / N_b, e_b = E_b / N_b, i_b = I_b / N_b,$$

$$r_b = R_b / N_b, w_a = W_a / N_a, z_b = Z_b / N_a$$

then the reduced equations become

To find the steady states, we set the right hand side of eqs. (15)-(24) to zero. Thus, the positive steady state solution is defined by

$$(s_a^*, e_a^*, i_a^*, r_a^*, w_a^*, s_b^*, e_b^*, i_b^*, r_b^*, w_b^*)$$

where

$$s_a^* = \frac{a\mu_h N}{(c_1 + \mu_h)N_a}, \quad (25)$$

$$e_a^* = \frac{ac_1\mu_h N}{(c_1 + \mu_h)(c_2 + \mu_h)N_a}, \quad (26)$$

$$i_a^* = \frac{ac_1c_2\mu_h N}{(c_1 + \mu_h)(c_2 + \mu_h)(c_3 + \mu_h)N_a}, \quad (27)$$

$$r_a^* = \frac{((\xi + c_6 + \mu_h) (-a\mu_h (c_7 (c_2 + \mu_h) (\xi + \mu_h) + c_1 (c_2 (-\xi + c_7) + c_7 (c_3 + \mu_h)))) N + c_7 (c_1 + \mu_h) (c_2 + \mu_h) (\xi + \mu_h) N_a))}{((\xi + \mu_h) (c_2 + \mu_h) (\xi + \mu_h) ((\xi + c_6 + \mu_h) (c_7 + \mu_h) + c_4 (\xi + c_7 + \mu_h)) N_a)}, \quad (28)$$

$$w_a^* = \frac{(c_4 (-a \mu_h (c_7 (c_2 + \mu_h) (c_3 + \mu_h) + c_1 (c_2 (-c_3 + c_7) + c_7 (c_3 + \mu_h)))) N + c_7 (c_1 + \mu_h) (c_2 + \mu_h) (c_3 + \mu_h) N_a))}{((c_1 + \mu_h) (c_2 + \mu_h) (c_3 + \mu_h) ((c_5 + c_6 + \mu_h) (c_7 + \mu_h) + c_4 (c_5 + c_7 + \mu_h)) N_a)}, \quad (29)$$

$$s_b^* = \frac{(1-a)\mu_h N}{(e_1 + \mu_h) N_b}, \quad (30)$$

$$e_b^* = \frac{(1-a)e_1 \mu_h N}{(e_1 + \mu_h)(e_2 + \mu_h) N_b}, \quad (31)$$

$$i_b^* = \frac{(1-a)e_1 e_2 \mu_h N}{(e_1 + \mu_h)(e_2 + \mu_h)(e_3 + \mu_h) N_a}, \quad (32)$$

$$r_b^* = \frac{((e_5 + e_6 + m_k) ((-1+a)m_k (e_7 (e_2 + m_k) (e_3 + m_k) + e_1 (e_2 (-e_3 + e_7) + e_7 (e_3 + m_k)))) N + e_7 (e_1 + m_k) (e_2 + m_k) (e_3 + m_k) N_b))}{((e_1 + m_k) (e_2 + m_k) (e_3 + m_k) ((e_5 + e_6 + m_k) (e_7 + m_k) + e_4 (e_5 + e_7 + m_k)) N_b)}, \quad (33)$$

$$w_b^* = \frac{((1-a) e_1 e_2 e_4 (e_3 - e_7) \mu_h N + e_4 e_7 (e_1 + \mu_h) (e_2 + \mu_h) (e_3 + \mu_h) N_b)}{((e_1 + \mu_h) (e_2 + \mu_h) (e_3 + \mu_h) ((e_5 + e_6 + \mu_h) (e_7 + \mu_h) + e_4 (e_5 + e_7 + \mu_h)) N_b)}, \quad (34)$$

III. ANALYTICAL SOLUTIONS

The local stable of each steady state is determined by the sign of eigenvalues for each steady state. If all eigenvalues have negative real parts, then that steady state is local stable [11]. The eigenvalues are the results of the following characteristic equation:

$$\det(J_e - \lambda I_{10}) = 0 \quad (35)$$

where I_{10} is the identity matrix dimension 10×10 and J_e is the *Jacobian* matrix of the positive steady state. From solving (17), the eigen values are given by

$$\begin{aligned} \lambda_1 &= -c_1 - \mu_h, \\ \lambda_2 &= -c_2 - \mu_h, \\ \lambda_3 &= -c_3 - \mu_h, \\ \lambda_{4,5} &= \frac{1}{2}(-c_4 - c_5 - c_6 - c_7 - 2\mu_h) \\ &\pm \sqrt{c_4^2 + (c_5 + c_6 - c_7)^2 - 2c_4(c_5 - c_6 + c_7)} \\ \lambda_6 &= -e_1 - \mu_h, \\ \lambda_7 &= -e_2 - \mu_h, \\ \lambda_8 &= -e_3 - \mu_h, \\ \lambda_{9,10} &= \frac{1}{2}(-e_4 - e_5 - e_6 - e_7 - 2\mu_h) \\ &\pm \sqrt{e_4^2 + (e_5 + e_6 - e_7)^2 - 2e_4(e_5 - e_6 + e_7)} \end{aligned} \quad (36)$$

We can see that all eigenvalues have negative real parts for

$R_0^a > 1$ and $R_0^b > 1$ where

$$R_0^a = \frac{c_4(c_4 + 2c_6) + (c_5 + c_6)^2 + c_7^2}{2c_4c_5 + 2(c_4 + c_5 + c_6)c_7}$$

and

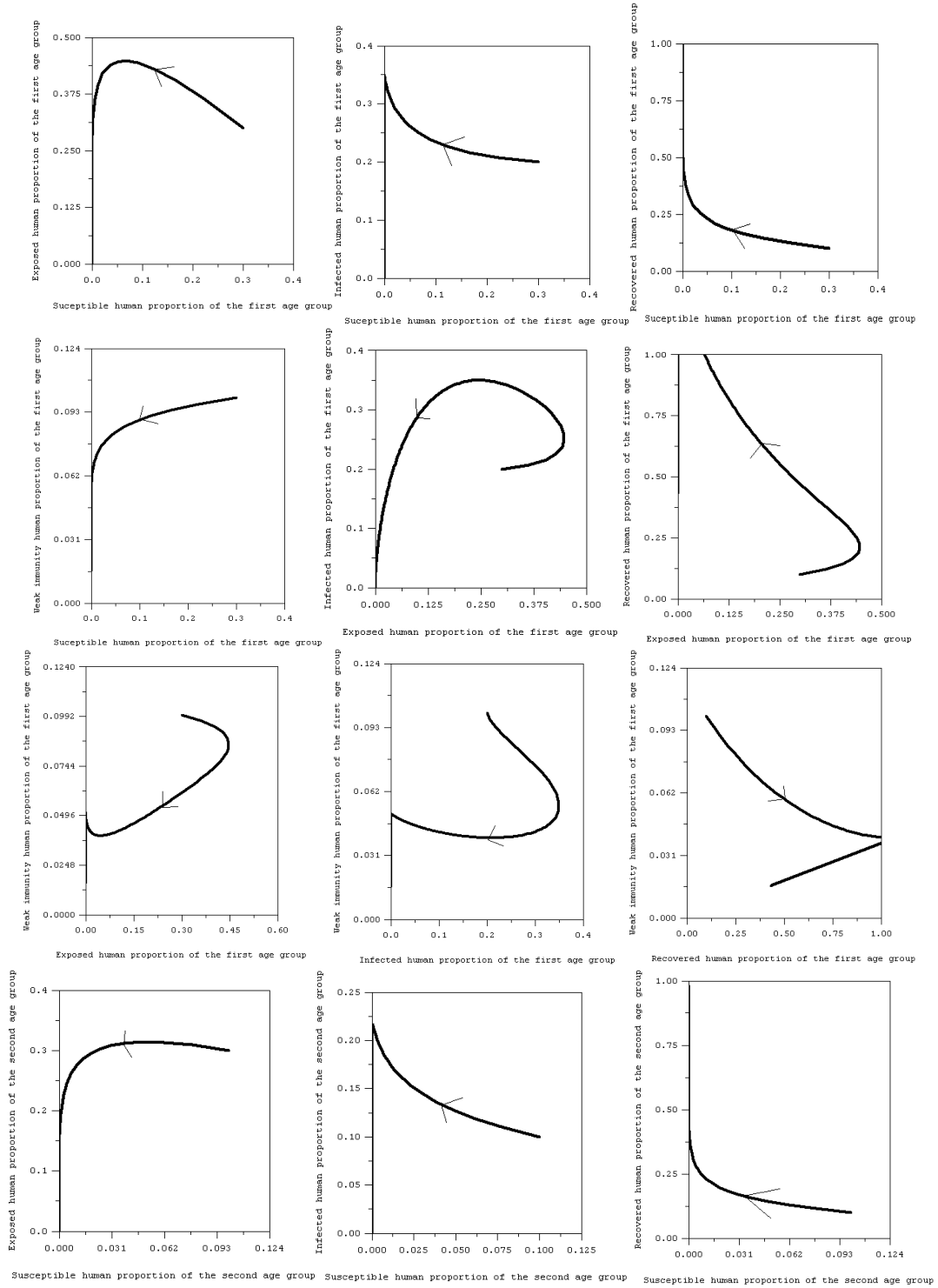
$$R_0^b = \frac{e_4(e_4 + 2e_6) + (e_5 + e_6)^2 + e_7^2}{2c_4c_5 + 2(c_4 + c_5 + c_6)e_7}$$

Therefore, the positive state is local stable for $R_0^a > 1$ and $R_0^b > 1$.

IV. NUMERICAL SOLUTIONS

The numerical solutions of our model are shown with the parameters defined as follows: $\mu_h = 1/(365*70)$; $a = 0.4$, $c_1 = 0.6$, $c_2 = 1/10$, $c_3 = 1/14$, $c_4 = 1/(365*2)$, $c_5 = 1/(365*2)$, $c_6 = 1/14$, $c_7 = 1/14$, $e_1 = 0.6$, $e_2 = 1/10$; $e_3 = 1/14$; $e_4 = 1/365$, $e_5 = 1/365$, $e_6 = 1/14$, $e_7 = 1/10$. Each parameters are obtained from simulation. $R_0^a = 1.01$ and $R_0^b = 1.03$. Steady state solution is given by (0.000043, 0.00026, 0.00037, 0.433348, 0.015999, 0.000065, 0.000391, 0.000547, 0.980208, 0.0184349)

From our simulation, we can see that the solutions converge to the non-trivial solution for $R_0^a > 1$ and $R_0^b > 1$ corresponding to the analytical solutions.



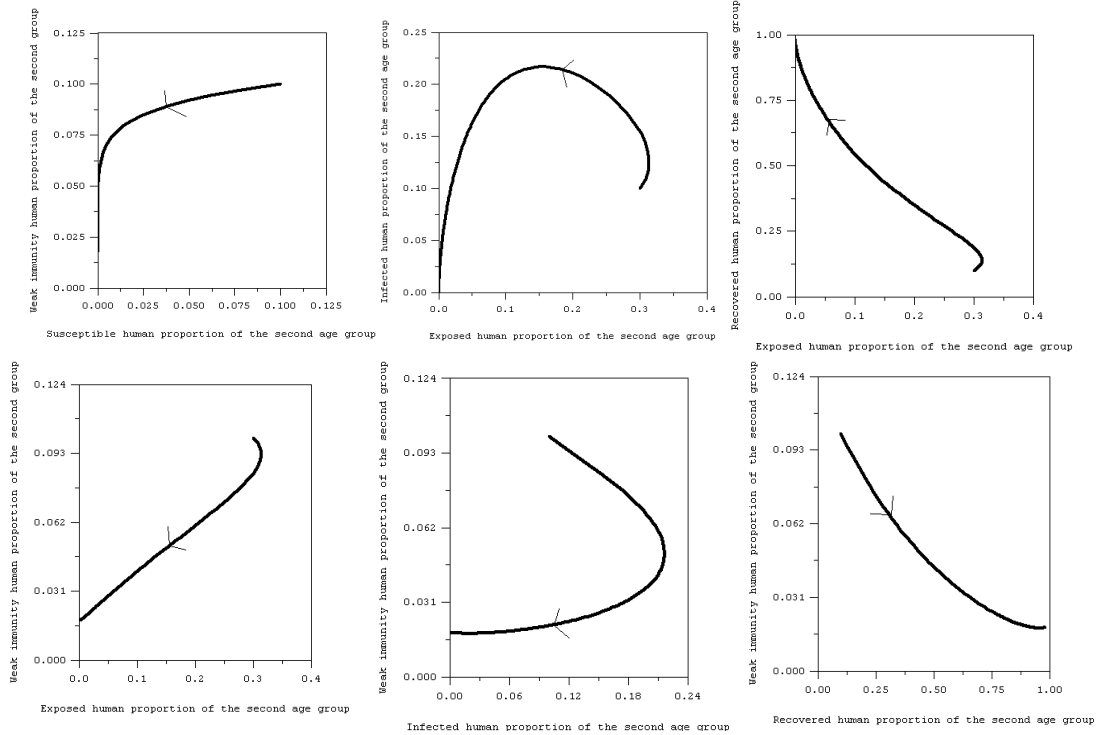


Fig. 3 Numerical solutions of our dynamical equations

V. DISCUSSION AND CONCLUSION

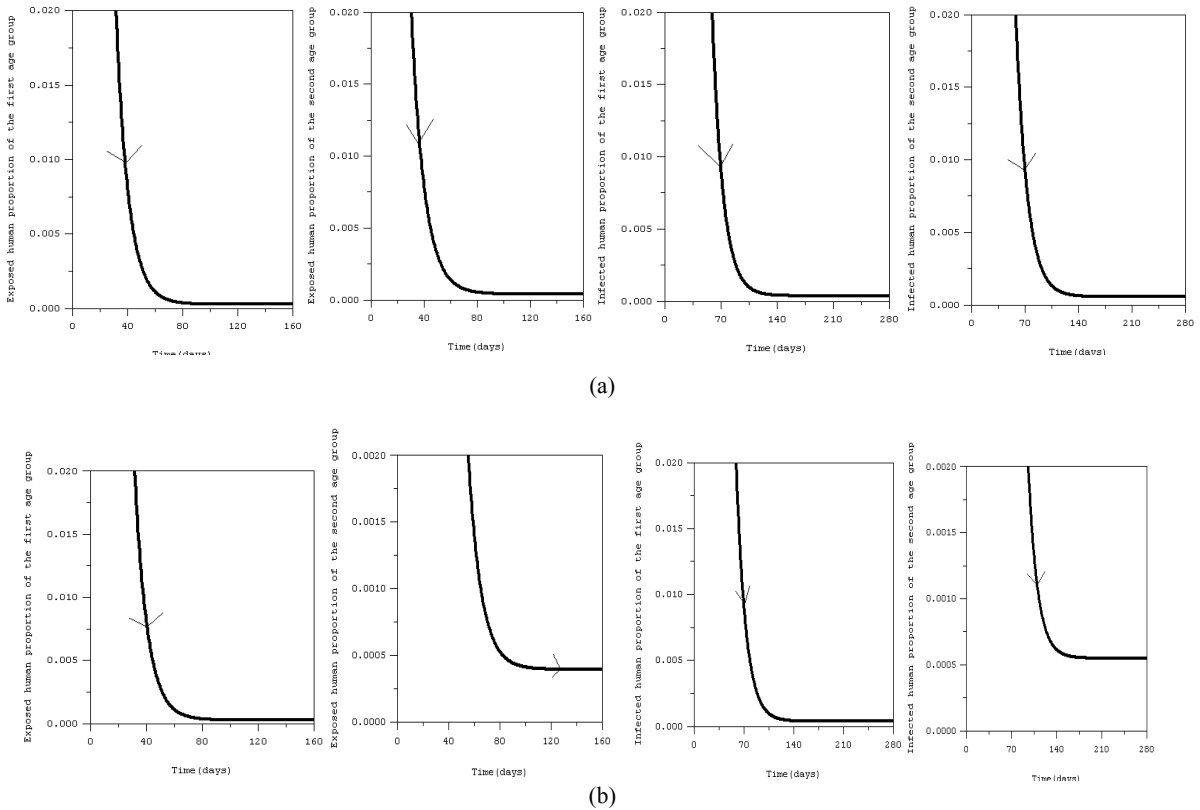


Fig. 4 Time series solutions for the different basic reproductive number (a) $R_0^a = 1.01$ and $R_0^b = 1.03$ (b) $R_0^a = 1.1$ and $R_0^b = 1.2$

From our analysis, we found the condition for local stability. We found that if $R_0^a > 1$ and $R_0^b > 1$ then the positive steady state is local stable, where

$$R_0^a = \frac{c_4(c_4 + 2c_6) + (c_5 + c_6)^2 + c_7^2}{2c_4c_5 + 2(c_4 + c_5 + c_6)c_7}$$

and
$$R_0^b = \frac{e_4(e_4 + 2e_6) + (e_5 + e_6)^2 + e_7^2}{2c_4c_5 + 2(c_4 + c_5 + c_6)e_7}.$$

We defined R_0^a and R_0^b as the basic reproductive number of this disease. It represents the average number of secondary cases reproduced from the primary cases [12]-[15]. From Fig. 4, we can see that the different basic reproductive number can produce the different outburst time and highest cases. The results of this study should introduce the way for reducing the transmission of this disease.

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