
Research article

Global analysis of a diffusive Cholera model with multiple transmission pathways, general incidence and incomplete immunity in a heterogeneous environment

Shengfu Wang and Linfei Nie*

College of Mathematics and System Science, Xinjiang University, Urumqi 830017, China

* **Correspondence:** Email: lfnie@163.com, nielinfei@xju.edu.cn.

Abstract: With the consideration of the complexity of the transmission of Cholera, a partially degenerated reaction-diffusion model with multiple transmission pathways, incorporating the spatial heterogeneity, general incidence, incomplete immunity, and Holling type II treatment was proposed. First, the existence, boundedness, uniqueness, and global attractiveness of solutions for this model were investigated. Second, one obtained the threshold condition \mathcal{R}_0 and gave its expression, which described global asymptotic stability of disease-free steady state when $\mathcal{R}_0 < 1$, as well as the maximum treatment rate as zero. Further, we obtained the disease was uniformly persistent when $\mathcal{R}_0 > 1$. Moreover, one used the mortality due to disease as a branching parameter for the steady state, and the results showed that the model undergoes a forward bifurcation at \mathcal{R}_0 and completely excludes the presence of endemic steady state when $\mathcal{R}_0 < 1$. Finally, the theoretical results were explained through examples of numerical simulations.

Keywords: reaction-diffusion model; vaccination; multiple routes of transmission; bifurcation analysis; Holling type II treatment

1. Introduction

Cholera is an emergency enteric epidemic induced by *Vibrio cholerae* (*V. cholerae*), and transmission of this disease is greatly compounded by interactions among host, pathogen, and environment. More specially, it can be transmitted by drinking or eating unpasteurized food or water infected with *V. cholerae*, by touching people with Cholera, hands and objects contaminated with the carrier's excreta, and by eating food contaminated with flies [1, 2]. Symptoms such as vomiting, leg cramps, and diarrhea can occur within 12 hours to five days after infection. World Health Organization (WHO) assessed that between 1.3 and 4 million incidences of Cholera occur and between 21,000 and 143,000 die annually, with children in Africa and Southeast Asia being the most impacted [3]. Cholera can also break out

commonly in countries with weak infrastructure and health systems, such as, Yemen, where 1,115,378 cases of suspected Cholera as well as 2310 deaths were reported between April 2017 and July 2018 [4]. The treatment methods for controlling Cholera include vaccines, rehydration therapy, and antibiotics. At present, vaccines are extensively used in certain areas; for example, Haiti successfully controlled the Cholera outbreak with the vaccine in 2020 [5].

Dynamical models have contributed to an essential role in gaining insight into the transmission mechanism, development process, and transmission pattern of Cholera, providing a contribution to the defense and management of strategic diseases. To date, a large number of scholars have devoted themselves to the study of Cholera (see, e.g., [6–11]), and the majority of models are characterized by ordinary/partial differential equations. The research contents include the nonnegative and boundedness of solutions, the persistence and extinction of this disease, bifurcation and chaotic phenomena, etc. In particular, Teytsa et al. [12] established the influence of phage bacterial invasion and optimal control on indirect transmission of Cholera disease by demonstrating that the release of lytic phages dramatically reduced the transmission of disease. Vaccines have always performed an active role in controlling and eradicating diseases, and this is also true for Cholera. For instance, Lin et al. [13] presented a Cholera model with high infectivity, low infectivity, and incomplete immunity, characterized the global dynamics of the equilibria, and simulated the Cholera epidemic in Haiti. In addition, there have also been numerous proposals for Cholera models with age structure [14], patch model [7], multiple disease stages [15], and so on. The relevant researches are still continuing.

As we all know, the propagation of Cholera is closely linked to numerous factors, for example, environmental sanitation, water and food resources, personal habits, and spatial heterogeneity. Lately, several reaction-diffusion models with environmental heterogeneity were developed to explore effective control strategies to eliminate this disease [16–18]. Specifically, Wang et al. [19] introduced a model in a closed environment and conducted a bifurcation analysis of the steady-state solution, which showed that spatial heterogeneity of model parameters can generate backward bifurcation. Avila-Vales et al. [20] proposed a *SIR* model with saturation incidence and Holling type II treatment, and theoretical results suggest that heterogeneity in transmission rates produces bifurcation, which leads to disease persistence. In [21–23], authors presented some reaction-diffusion models and attained \mathcal{R}_0 , which examined the presence as well as global stability of steady states. Wang et al. [24] established a reaction-convection-diffusion model based on the high infectivity of bacteria, and revealed that ignoring high infectivity underestimates the risk of illness propagation. Wang et al. [25] also developed a Cholera transmission model with high bacterial infectivity and different diffusion rates, assuming different transmission rates for susceptible and infected individuals, which showed that by controlling the mobility of susceptible individuals, the illness would be eradicated to some extent.

Motivated by the previous works, in this article, a reaction-diffusion model of Cholera transmission with horizontal/environmental propagation as well as general incidence is proposed, where the incomplete immunity, Holling II treatment rates, different diffusion rates, and environmental viruses are also introduced. The remaining sections of this article are structured below: In Sections 2 and 3, the model is established and the well-posedness of the model is analyzed. The basic reproduction number is presented and the global stability of disease-free steady state and the persistence of disease is analyzed in Section 4. The positive steady state of the model is investigated in Section 5 from the branching theory point of view. Some numerical simulations and a short summary are given in Sections 6 and 7, respectively.

2. Mathematical model

Following the pattern of Cholera transmission, the population of a given area is divided as: susceptible, vaccinated, infected, and recovered individuals are represented by $S(x, t)$, $V(x, t)$, $I(x, t)$, and $R(x, t)$, respectively. Further, the concentration of pathogen particles is represented by $W(x, t)$. The corresponding flow chart for Cholera propagation is shown in Figure 1. Based on the variability of Cholera spreading routes and the restricted diffusion of *V. cholerae* in the environment, the partially degenerate reaction-diffusion model is described by

$$\begin{cases} \frac{\partial S}{\partial t} = d_1 \Delta S + \Lambda(x) - (\mu(x) + \rho(x))S - F(x, S, I) - G(x, S, W) + \theta(x)V, \\ \frac{\partial V}{\partial t} = d_2 \Delta V + \rho(x)S - \sigma F(x, V, I) - \sigma G(x, V, W) - (\mu(x) + \theta(x))V, \\ \frac{\partial I}{\partial t} = d_3 \Delta I + F(x, S, I) + G(x, S, W) + \sigma(F(x, V, I) + G(x, V, W)) \\ \quad - (\mu(x) + d(x) + r(x))I - \frac{\gamma(x)I}{1 + a(x)I}, \\ \frac{\partial W}{\partial t} = \alpha(x)I - \xi(x)W, \end{cases} \quad (2.1)$$

and

$$\frac{\partial R}{\partial t} = d_4 \Delta R + r(x)I + \frac{\gamma(x)I}{1 + a(x)I} - \mu(x)R,$$

subject to the boundary conditions

$$\frac{\partial S}{\partial n} = \frac{\partial V}{\partial n} = \frac{\partial I}{\partial n} = \frac{\partial W}{\partial n} = \frac{\partial R}{\partial n} = 0, \quad t > 0, \quad x \in \partial \mathbb{D}, \quad (2.2)$$

and initial conditions $S(0, x) = S_0(x)$, $V(0, x) = V_0(x)$, $I(0, x) = I_0(x)$, $V(0, x) = V_0(x)$, $R(0, x) = R_0(x)$, $W(0, x) = W_0(x)$, $x \in \mathbb{D}$, where \mathbb{D} is a connected, bounded subset of \mathbb{R}^n with smooth boundary $\partial \mathbb{D}$. The means of other model parameters are as: $d_1, d_2, d_3, d_4 > 0$ stand for the diffusion rates measuring the movement for susceptible, vaccinated, infected, and recovered individuals, respectively; $\Lambda(x)$, $\mu(x)$, $d(x)$, $\rho(x)$, $\theta(x)$ stand for the population replenishment rate, the natural mortality rate, the disease-related mortality rate, the vaccination rate, and the rate of loss of immunization, respectively; $r(x)$ denotes the natural recovery rate; $\alpha(x)$ represents the bacterial shedding rate of infected individuals and $\xi(x)$ stands for the decay rate of bacteria; $\gamma(x)/(1 + a(x))$ denotes the treatment function, where $\gamma(x)$ stands for the maximum treatment rate per individual per unit of time, and $a(x)$ represents the influence of delayed treatment in infected individuals; σ denotes the reduction of vaccine efficacy; $F(x, S, I)$ and $G(x, S, W)$ indicate general incidence functions responding to direct transmission from infected individuals to susceptible individuals and indirect transmission from environmental viruses to susceptible individuals, respectively; $F(x, V, I)$ and $G(x, V, W)$ correspond to transmission between vaccinated individuals and infected individuals and between vaccinated individuals and environmental viruses, respectively.

As model (2.1) does not contain the variable R , one can overlook the R -equation and restrict attention to the kinetic behavior of model (2.1). In light of the model's biological context, all parameters are hypothesized to be positive, continuous, and bounded on $\bar{\mathbb{D}}$. Moreover, let us hypothesize that the functions F and G fulfill the under mentioned cases.

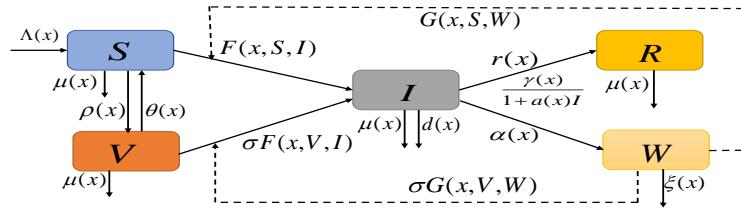


Figure 1. A dynamic Cholera propagation graph of model (2.1).

- (H₁) For $x \in \overline{\mathbb{D}}$ and $S, V, I \geq 0$, $F(x, S, 0) = F(x, 0, I) = 0$, $F(x, V, 0) = F(x, 0, I) = 0$ and $\partial F(x, S, I)/\partial I > 0$, $\partial F(x, V, I)/\partial I > 0$, $\partial^2 F(x, S, I)/\partial I^2 \leq 0$, $\partial^2 F(x, V, I)/\partial I^2 \leq 0$.
- (H₂) For $x \in \overline{\mathbb{D}}$ and $S, V, W \geq 0$, $G(x, S, 0) = G(x, 0, W) = 0$, $G(x, V, 0) = G(x, 0, W) = 0$; $\partial G(x, S, W)/\partial W > 0$, $\partial G(x, V, W)/\partial W > 0$, and $\partial^2 G(x, S, W)/\partial W^2 \leq 0$, $\partial^2 G(x, V, W)/\partial W^2 \leq 0$.
- (H₃) There are Hölder continuous functions $\beta_i : \mathbb{D} \rightarrow \mathbb{R}_+$ that satisfy $F(x, y, I) \leq \beta_1(x)yI$, $G(x, y, W) \leq \beta_2(x)yW$, $y \in \{S, V\}$, for $x \in \mathbb{D}$, $S, I, V, W \geq 0$.

Remark 2.1. Some frequently used incidences satisfy (H₁) and (H₂), such as the bilinear incidence rates $F(x, S, I) = \beta_1(x)SI$, $G(x, S, W) = \beta_2(x)SW$ (see [7, 15]); the saturated incidence rates $F(x, S, I) = \beta_1(x)SI/(\kappa_1(x) + I)$, $G(x, S, W) = \beta_2(x)SP/(\kappa_2(x) + P)$ (see [26]), where $\beta_i(x), \kappa_i(x) > 0$, $i = 1, 2$. The condition (H₃) is given to better prove the fitness of solution. At the same time, we also find that the above common incidence also satisfies the condition (H₃).

3. Well-posedness

Let $\mathbb{X} := C(\overline{\mathbb{D}}, \mathbb{R}^4)$ be the Banach space, and define $\mathbb{X}^+ := C(\overline{\mathbb{D}}, \mathbb{R}_+^4)$. Set $\psi^+ := \max_{x \in \overline{\mathbb{D}}} \{\psi(x)\}$, $\psi^- := \min_{x \in \overline{\mathbb{D}}} \{\psi(x)\}$, where ψ represents any of $\Lambda, \mu, \rho, \theta, r, \gamma, a, \alpha, \xi$.

To do so, denote $\pi_1(x) = \rho(x) + \mu(x)$, $\pi_2(x) = \theta(x) + \mu(x)$, $\pi_3(x) = d(x) + r(x) + \mu(x)$, and $\pi_4(x) = \xi(x)$, and let $\Gamma_i(t) : C(\overline{\mathbb{D}}, \mathbb{R}) \rightarrow C(\overline{\mathbb{D}}, \mathbb{R})$ ($i = 1, 2, 3$) be the C_0 semigroup related to $d_i \Delta - \pi_i(x)$ satisfying the Neumann boundary condition. Hence,

$$(\Gamma_i(t)\phi)(x) = \int_{\Omega} T_i(x, t, y) \phi(y) dy, \quad \forall t > 0, \phi \in C(\overline{\mathbb{D}}, \mathbb{R}), i = 1, 2, 3,$$

where $T_i(x, t, y)$ denotes the Green function related to $d_i \Delta - \pi_i(x)$ satisfying (2.2). Further, let $(\Gamma_4(t)\phi)(x) = e^{-\pi_4(x)t} \phi(x)$. Thus, $\Gamma(t) := \text{diag}\{\Gamma_1(t), \Gamma_2(t), \Gamma_3(t), \Gamma_4(t)\} : \mathbb{X} \rightarrow \mathbb{X}$, $t \geq 0$, which formulates a strongly continuous semigroup [27].

For every $\phi = (\phi_1, \phi_2, \phi_3, \phi_4)^T \in \mathbb{X}^+$, define $Z = (Z_1, Z_2, Z_3, Z_4)^T : \mathbb{X}^+ \rightarrow \mathbb{X}$ by

$$Z_1(\phi)(x) = \Lambda(x) - F(x, \phi_1, \phi_3) - G(x, \phi_1, \phi_4) + \theta(x)\phi_2, \quad Z_4(\phi)(x) = \alpha(x)\phi_3,$$

$$Z_2(\phi)(x) = \rho(x)\phi_1 - \sigma F(x, \phi_2, \phi_3) - \sigma G(x, \phi_2, \phi_4),$$

$$Z_3(\phi)(x) = F(x, \phi_1, \phi_3) + G(x, \phi_1, \phi_4) + \sigma F(x, \phi_2, \phi_3) + \sigma G(x, \phi_2, \phi_4) - \frac{\gamma(x)\phi_3}{1 + a(x)\phi_3},$$

where T denotes the transposition. Hence, model (2.1) can be reformulated as

$$u(t) = \Gamma(t)\phi + \int_0^t \Gamma(t-s)Z(u(s)) ds. \quad (3.1)$$

In the results below, the local solutions of model (2.1) with (2.2) on \mathbb{X}^+ are involved.

Lemma 3.1. *For $\phi \in \mathbb{X}^+$, model (2.1) possesses a unique solution $u(\cdot, t, \phi) := (S(\cdot, t), V(\cdot, t), I(\cdot, t), W(\cdot, t))$ on $[0, \tau_{\max}]$ with $u(\cdot, 0, \phi) = \phi$, where $\tau_{\max} \leq \infty$. Moreover, $u(\cdot, t, \phi) \in \mathbb{X}^+$, $0 \leq t < \tau_{\max}$.*

Proof. For $h \geq 0$, one obtains

$$\begin{aligned} \phi + h\Gamma(\phi) &= \begin{pmatrix} \phi_1 + h[\Lambda(x) - F(x, \phi_1, \phi_3) - G(x, \phi_1, \phi_4) + \theta(x)\phi_2] \\ \phi_2 + h[\rho(x)\phi_1 - \sigma F(x, \phi_2, \phi_3) - \sigma G(x, \phi_2, \phi_4)] \\ \phi_3 + h[F(x, \phi_1, \phi_3) + G(x, \phi_1, \phi_4) + \sigma F(x, \phi_2, \phi_3) + \sigma G(x, \phi_2, \phi_4) - \frac{\gamma(x)\phi_3}{1+a(x)\phi_3}] \\ \phi_4 + h\alpha(x)\phi_3 \end{pmatrix} \\ &\geq \begin{pmatrix} \phi_1 - h[F(x, \phi_1, \phi_3) + G(x, \phi_1, \phi_4) - \theta(x)\phi_2] \\ \phi_2 - h[\sigma F(x, \phi_2, \phi_3) + \sigma G(x, \phi_2, \phi_4)] \\ \phi_3 - h\frac{\gamma(x)\phi_3}{1+a(x)\phi_3} \\ \phi_4 \end{pmatrix}, \end{aligned}$$

which means for $\phi \in \mathbb{X}^+$, $\lim_{h \rightarrow 0^+} \text{dist}(\phi + h\Gamma(\phi), \mathbb{X}^+) = 0$. Based on Ref. [28, Corollary 4], model (2.1) is a unique mild solution $(S(x, t), V(x, t), I(x, t), W(x, t))$ on $[0, \tau_{\max}]$, where $\tau_{\max} \leq \infty$. \square

Consider the model as below

$$\frac{\partial \omega}{\partial t} = d\Delta\omega + b(x) - c(x)\omega, \quad x \in \mathbb{D}, \quad t > 0; \quad \frac{\partial \omega}{\partial n} = 0, \quad x \in \partial\mathbb{D}, \quad (3.2)$$

where $d > 0$, $b(x) > 0$, and $c(x) > 0$ are continuous.

Lemma 3.2 (Lemma 1 in Ref. [29]). *Model (3.2) possesses a globally asymptotically stable steady state $\hat{\omega}(x)$ in $C(\overline{\mathbb{D}}, \mathbb{R}_+)$. Further, if $b(x) \equiv b$, $c(x) \equiv c$, $\forall x \in \overline{\mathbb{D}}$, then $\hat{\omega}(x) = b/c$.*

Next, one proves that the local solution of model (2.1) can be expanded to the global solution, i.e., $\tau_{\max} = \infty$.

Lemma 3.3. *For every $\phi \in \mathbb{X}^+$, model (2.1) has a unique solution $u(x, t)$ with $u(x, 0, \phi) = \phi$ for $[0, \infty)$. Moreover, the model generates a semiflow $\Phi(t)$ as ultimately bounded.*

Proof. In terms of the first two equations of the model (2.1), one can readily derive that

$$\begin{cases} \frac{\partial S}{\partial t} \leq d_1\Delta S + \Lambda^+ - (\mu^- + \rho^-)S + \theta^+V, & x \in \mathbb{D}, \quad t > 0, \\ \frac{\partial V}{\partial t} \leq d_2\Delta V + \rho^+S - (\mu^- + \theta^-)V, & x \in \mathbb{D}, \quad t > 0, \\ \frac{\partial S}{\partial n} = \frac{\partial V}{\partial n} = 0, & x \in \partial\mathbb{D}, \quad t > 0. \end{cases}$$

Hence,

$$\begin{aligned} \limsup_{t \rightarrow \infty} S(x, t) &\leq \frac{\Lambda^+(\mu^- + \theta^-)}{(\mu^- + \rho^-)(\mu^- + \theta^-) - \theta^+\rho^+} := N_1, \\ \limsup_{t \rightarrow \infty} V(x, t) &\leq \frac{\Lambda^+\rho^+(\mu^- + \theta^-)}{(\mu^- + \theta^-)((\mu^- + \rho^-)(\mu^- + \theta^-) - \theta^+\rho^+)} := N_2, \quad \text{uniformly in } x \in \overline{\mathbb{D}}, \end{aligned} \quad (3.3)$$

which implies $\|S(x, t)\| \leq \mathcal{M}_1$, $\|V(x, t)\| \leq \mathcal{M}_2$, for $\mathcal{M}_1, \mathcal{M}_2 > 0$ and $0 \leq t < \infty$. Hence, one gets that $S(x, t)$ and $V(x, t)$ are ultimately bounded. Adding the three previous equations of (2.1) and integrating with respect to \mathbb{D} gives

$$\frac{\partial}{\partial t} \int_{\mathbb{D}} (S(x, t) + V(x, t) + I(x, t)) dx \leq \Lambda^+ |\mathbb{D}| - \mu^- \int_{\mathbb{D}} (S(x, t) + V(x, t) + I(x, t)) dx,$$

where $|\mathbb{D}|$ denotes the measurement of region \mathbb{D} . It follows that

$$\limsup_{t \rightarrow \infty} (\|S(x, t)\|_1 + \|V(x, t)\|_1 + \|I(x, t)\|_1) \leq \mathcal{M}_{11},$$

with $\mathcal{M}_{11} = \Lambda^+ |\mathbb{D}| / \mu^-$. In a similar way, the W -equation satisfies

$$\frac{\partial}{\partial t} \int_{\mathbb{D}} W(x, t) dx \leq \alpha^+ \mathcal{M}_{11} - \xi^- \int_{\mathbb{D}} W(x, t) dx.$$

Thus, one gets

$$\limsup_{t \rightarrow \infty} \|W(x, t)\|_1 \leq \mathcal{M}_{12}, \text{ with } \mathcal{M}_{12} = \frac{\alpha^+ \mathcal{M}_{11}}{\xi^-}.$$

In short, there exists a number $\mathcal{M}_3 > 0$,

$$\limsup_{t \rightarrow \infty} (\|S(x, t)\|_1 + \|V(x, t)\|_1 + \|I(x, t)\|_1 + \|W(x, t)\|_1) \leq \mathcal{M}_3,$$

Next, let us verify the solution (I, W) of model (2.1) as ultimately bounded. Motivated by [30, Lemma 2.4], for $T > 0$, one needs to justify

$$\limsup_{t \rightarrow \infty} (\|I(x, t)\|_{2^k} + \|W(x, t)\|_{2^k}) \leq \mathcal{M}_{2^k}, \quad \forall t > T, \quad (3.4)$$

where $\mathcal{M}_{2^k} > 0$ is a constant.

It immediately follows that for $k = 0$, (3.4) holds. Assuming that (3.4) is valid for $k - 1$, i.e.,

$$\limsup_{t \rightarrow \infty} (\|I(x, t)\|_{2^{k-1}} + \|W(x, t)\|_{2^{k-1}}) \leq \mathcal{M}_{2^{k-1}}, \quad \text{for } \mathcal{M}_{2^{k-1}} > 0. \quad (3.5)$$

The I -equation of model (2.1) is multiplied by $I^{2^{k-1}}$ and integrated over \mathbb{D} to derive

$$\begin{aligned} & \frac{1}{2^k} \frac{\partial}{\partial t} \int_{\mathbb{D}} I^{2^k} dx \\ & \leq d_3 \int_{\mathbb{D}} I^{2^{k-1}} \Delta I dx + \int_{\mathbb{D}} F(x, S, I) I^{2^{k-1}} dx + \int_{\mathbb{D}} G(x, S, W) I^{2^{k-1}} dx - \int_{\mathbb{D}} \frac{\gamma(x) I^{2^k}}{1 + a(x) I} dx \\ & \quad + \int_{\mathbb{D}} \sigma F(x, V, I) I^{2^{k-1}} dx + \int_{\mathbb{D}} \sigma G(x, V, W) I^{2^{k-1}} dx - \int_{\mathbb{D}} (\mu(x) + d(x)) I^{2^k} dx \\ & \leq d_3 \int_{\mathbb{D}} I^{2^{k-1}} \Delta I dx + \int_{\mathbb{D}} \beta_1(x) S I^{2^k} dx + \int_{\mathbb{D}} \beta_2(x) S W I^{2^{k-1}} dx + \int_{\mathbb{D}} \sigma \beta_1(x) V I^{2^k} dx \\ & \quad + \int_{\mathbb{D}} \sigma \beta_2(x) V W I^{2^{k-1}} dx - \int_{\mathbb{D}} (\mu(x) + d(x) + r(x)) I^{2^k} dx. \end{aligned} \quad (3.6)$$

Recall that

$$\begin{aligned} d_3 \int_{\mathbb{D}} I^{2^{k-1}} \Delta I dx &\leq -d_3 \int_{\mathbb{D}} \nabla I \cdot \nabla I^{2^{k-1}} dx = -(2^k - 1)d_3 \int_{\mathbb{D}} (\nabla I \cdot \nabla I) I^{2^{k-2}} dx \\ &= -\frac{2^k - 1}{2^{2k-2}} d_3 \int_{\mathbb{D}} |\nabla I^{2^{k-1}}|^2 dx. \end{aligned}$$

Hence, inequality (3.6) becomes

$$\begin{aligned} \frac{1}{2^k} \frac{\partial}{\partial t} \int_{\mathbb{D}} I^{2^k} dx &\leq -L_k \int_{\mathbb{D}} |\Delta I^{2^{k-1}}|^2 dx + \int_{\mathbb{D}} (\beta_1(x) S I^{2^k} + \beta_2(x) S W I^{2^{k-1}}) dx + \int_{\mathbb{D}} \sigma \beta_1(x) V I^{2^k} dx \\ &\quad + \int_{\mathbb{D}} \sigma \beta_2(x) V W I^{2^{k-1}} dx - \int_{\mathbb{D}} (r(x) + \mu(x) + d(x)) I^{2^k} dx, \end{aligned} \quad (3.7)$$

where $L_k = (2^k - 1)/(2^{2k-2})$. Due to $\limsup_{t \rightarrow \infty} \|S(x, t)\| \leq \mathcal{M}_1$, $\limsup_{t \rightarrow \infty} \|V(x, t)\| \leq \mathcal{M}_2$, there is $t_0 > 0$ satisfying when $t \geq t_0$, and one has

$$\begin{aligned} \int_{\mathbb{D}} \beta_1 S I^{2^k} dx &\leq \beta_1^+ (\mathcal{M}_1 + 1) \int_{\mathbb{D}} I^{2^k} dx, \quad \int_{\mathbb{D}} \beta_2 S W I^{2^{k-1}} dx \leq \beta_2^+ (\mathcal{M}_1 + 1) \int_{\mathbb{D}} W I^{2^{k-1}} dx, \\ \int_{\mathbb{D}} \sigma \beta_1 V I^{2^k} dx &\leq \sigma \beta_1^+ (\mathcal{M}_2 + 1) \int_{\mathbb{D}} I^{2^k} dx, \quad \int_{\mathbb{D}} \sigma \beta_2 V W I^{2^{k-1}} dx \leq \sigma \beta_2^+ (\mathcal{M}_2 + 1) \int_{\mathbb{D}} W I^{2^{k-1}} dx. \end{aligned} \quad (3.8)$$

By means of Young's inequality: $ab \leq \varepsilon a^p + \varepsilon^{-\frac{q}{p}} b^q$, where $a, b, \varepsilon > 0$, $1/p + 1/q = 1$. By setting $\varepsilon_1 = \xi^-/(4\beta_2^+(\mathcal{M}_0 + 1))$, $\mathcal{M}_0 = \max\{\mathcal{M}_1, \mathcal{M}_2\}$, $p = 2^k$, and $q = 2^k/(2^k - 1)$, we have

$$\int_{\mathbb{D}} W I^{2^{k-1}} dx \leq \frac{\xi^-}{4\beta_2^+(\mathcal{M}_1 + 1)} \int_{\mathbb{D}} W^{2^k} dx + C_{\varepsilon_1} \int_{\mathbb{D}} I^{2^k} dx, \text{ for } t \geq t_0, \text{ and } C_{\varepsilon_1} = \varepsilon_1^{-\frac{1}{2^{k-1}}}. \quad (3.9)$$

Thus, (3.7) is reorganized as:

$$\begin{aligned} \frac{1}{2^k} \frac{\partial}{\partial t} \int_{\mathbb{D}} I^{2^k} dx &\leq -L_k \int_{\mathbb{D}} |\nabla I^{2^{k-1}}|^2 dx + \beta_1^+ (\mathcal{M}_1 + 1) \int_{\mathbb{D}} I^{2^k} dx + \frac{\xi^-}{2} \int_{\mathbb{D}} W^{2^k} dx \\ &\quad + \beta_2^+ (\mathcal{M}_1 + 1) C_{\varepsilon_1} \int_{\mathbb{D}} I^{2^k} dx + \sigma \beta_1^+ (\mathcal{M}_2 + 1) \int_{\mathbb{D}} I^{2^k} dx \\ &\quad + \sigma \beta_2^+ (\mathcal{M}_2 + 1) \int_{\mathbb{D}} I^{2^k} dx + \sigma \beta_2^+ (\mathcal{M}_2 + 1) C_{\varepsilon_1} \int_{\mathbb{D}} I^{2^k} dx \\ &\leq -L_k \int_{\mathbb{D}} |\nabla I^{2^{k-1}}|^2 dx + C_k \int_{\mathbb{D}} I^{2^k} dx + \frac{\xi^-}{2} \int_{\mathbb{D}} W^{2^k} dx, \end{aligned} \quad (3.10)$$

where $C_k = \sigma \beta_1^+ (\mathcal{M}_2 + 1) + \sigma \beta_2^+ (\mathcal{M}_2 + 1) + \sigma \beta_2^+ (\mathcal{M}_2 + 1) C_{\varepsilon_1} + \beta_1^+ (\mathcal{M}_1 + 1) + \beta_2^+ (\mathcal{M}_2 + 1) C_{\varepsilon_1}$.

Similarly, multiplying the W -equation with $W^{2^{k-1}}$, one yields

$$\frac{1}{2^k} \frac{\partial}{\partial t} \int_{\mathbb{D}} W^{2^k} dx \leq \alpha^+ \int_{\mathbb{D}} I W^{2^{k-1}} dx - \xi^- \int_{\mathbb{D}} W^{2^k} dx. \quad (3.11)$$

By choosing $p = 2^k/(2^k - 1)$ and $q = 2^k$, we have

$$\int_{\mathbb{D}} I W^{2^{k-1}} dx \leq \frac{\xi^-}{4\alpha^+} \int_{\mathbb{D}} W^{2^k} dx + C_{\varepsilon_2} \int_{\mathbb{D}} I^{2^k} dx. \quad (3.12)$$

where $\varepsilon_2 = \xi^-/4\alpha^+$, $C_{\varepsilon_2} = \varepsilon_2^{1-2^k}$. Hence, (3.11) becomes

$$\frac{1}{2^k} \frac{\partial}{\partial t} \int_{\mathbb{D}} W^{2^k} dx \leq -\frac{3}{4} \xi^- \int_{\mathbb{D}} W^{2^k} dx + \alpha^+ C_{\varepsilon_2} \int_{\mathbb{D}} I^{2^k} dx. \quad (3.13)$$

Adding (3.10) and (3.13), one has

$$\frac{1}{2^k} \frac{\partial}{\partial t} \int_{\mathbb{D}} (I^{2^k} + W^{2^k}) dx \leq -L_k \int_{\mathbb{D}} |\nabla I^{2^{k-1}}|^2 dx + Q_k \int_{\mathbb{D}} I^{2^k} dx - \frac{\xi^-}{4} \int_{\mathbb{D}} W^{2^k} dx, \quad (3.14)$$

where $Q_k = \alpha^+ C_{\varepsilon_2} + C_k$. Applying the interpolation inequality,

$$\|\xi\|_2^2 \leq \|\nabla \xi\|_2^2 + C_\varepsilon \|\xi\|_1, \text{ where } \xi \in W^{1,2}(\mathbb{D}). \quad (3.15)$$

Let $\varepsilon_3 = L_k/(2Q_k)$, $\zeta = I^{2^{k-1}}$, then

$$-L_k \int_{\mathbb{D}} |\nabla I^{2^{k-1}}|^2 dx \leq -2Q_k \int_{\mathbb{D}} I^{2^k} dx + 2Q_k C_{\varepsilon_3} \left(\int_{\mathbb{D}} I^{2^{k-1}} dx \right)^2, \quad (3.16)$$

Therefore, inequality (3.14) becomes

$$\frac{1}{2^k} \frac{\partial}{\partial t} \int_{\mathbb{D}} (I^{2^k} + W^{2^k}) dx \leq -\varsigma_* \left(\int_{\mathbb{D}} I^{2^k} dx + \int_{\mathbb{D}} W^{2^k} dx \right) + 2Q_k C_{\varepsilon_3} \left(\int_{\mathbb{D}} I^{2^{k-1}} dx \right)^2, \quad (3.17)$$

where $\varsigma_* = \min\{Q_k, \xi^-/4\}$. It follows from (3.5) that $\limsup_{t \rightarrow \infty} \int_{\mathbb{D}} I^{2^{k-1}} dx \leq \mathcal{M}_{2^{k-1}}^{2^{k-1}}$, which means

$$\limsup_{t \rightarrow \infty} (\|I(x, t)\|_{2^k} + \|W(x, t)\|_{2^k}) \leq \mathcal{M}_{2^k}, \text{ with } \mathcal{M}_{2^k} = \sqrt[2^k]{\frac{2Q_k C_{\varepsilon_3}}{\varsigma_*}} \mathcal{M}_{2^k}.$$

Therefore, by the continuous embedding $L^q(\mathbb{D}) \hookrightarrow L^p(\mathbb{D})$, one has

$$\limsup_{t \rightarrow \infty} (\|I(x, t)\|_{L^p} + \|W(x, t)\|_{L^p}) \leq \mathcal{M}_p, \text{ for } q > p > 1,$$

where $\mathcal{M}_p > 0$ is a constant. The fractional power space is represented by Y_a ($0 \leq a \leq 1$). By Ref.[30, Lemma 2.4], one gets that $Y_a \subset C(\bar{\mathbb{D}})$ by choosing $p > n/2$ and $a \geq n/2p$. Thus, we can get

$$\limsup_{t \rightarrow \infty} \|I(x, t)\| \leq \mathcal{M}_\infty, \quad \limsup_{t \rightarrow \infty} \|W(x, t)\| \leq \frac{\alpha^+}{\xi^-} \mathcal{M}_\infty, \text{ where } \mathcal{M}_\infty > 0,$$

which demonstrates that Lemma 3.3 is valid. \square

Let

$$\mathbf{D} = \left\{ (S, V, I, W) \in \mathbb{X}^+ : S(x, t) \leq N_1, V(x, t) \leq N_2, I(x, t) \leq \mathcal{M}_\infty, W(x, t) \leq \frac{\alpha^+}{\xi^-} \mathcal{M}_\infty \right\},$$

then $\Phi(t)\phi \in \mathbf{D}$, $t \geq t_1$, for some $t_1 \geq 0$. In addition, in analogy to the approach in Ref. [31, Theorem 2.1], we learn for set $\mathcal{V} \subset \mathbb{X}^+$, $\Phi(t)\phi \in \mathbf{D}$, $t \geq t_2$, for $t_2 \geq 0$.

As the last equation of model (2.1) has no diffusion, the weak compactness of solution semiflow $\Phi(t)$ is hard to obtain, and we substitute the weak compactness with the asymptotic smoothness of the solution semiflow. At first, one defines the Kuratowski measure of noncompactness, $\tau(\cdot)$,

$$\tau(\mathcal{V}) := \inf\{r : \mathcal{V} \text{ has a finite cover of diameter } < r\},$$

for set $\mathcal{V} \subset \mathbb{X}^+$. It's convenient to deduce that \mathcal{V} is precompact if and only if $\kappa(\mathcal{V}) = 0$.

Denote $\mathbf{x} := (S, V, I)$, $\mathbf{y} := (W)$, and $\mathbf{g}(x, t, \mathbf{x}, \mathbf{y}) = -\alpha(x)I - \xi(x)W$. Taking the partial derivative of $\mathbf{g}(x, t, \mathbf{x}, \mathbf{y})$ relative to \mathbf{y} yields

$$\frac{\partial \mathbf{g}(x, t, \mathbf{x}, \mathbf{y})}{\partial \mathbf{y}} = -\xi(x) \leq -\xi^-.$$

Lemma 3.4. *In case there exists $q > 0$ satisfying*

$$\tau^T \frac{\partial \mathbf{g}(x, t, \mathbf{x}, \mathbf{y})}{\partial \mathbf{y}} \tau \leq -q \tau^T \tau, \quad \forall \tau \in \mathbb{R}, x \in \overline{\mathbb{D}}, W \in \mathbf{D},$$

then $\Phi(t)$ is κ -contracting, i.e., $\lim_{t \rightarrow \infty} \kappa(\Phi(t)\mathcal{V}) = 0$ for set $\mathcal{V} \subset \mathbb{X}^+$.

Proof. In a similar way to [32, Lemma 4.1], we can demonstrate that $\Phi(t)$ is asymptotically compact on \mathcal{V} , i.e., for $t_n \rightarrow \infty$ and any sequence $\phi_n \in \mathcal{V}$, a subsequence $t_{n_k} \rightarrow \infty$ and ϕ_{n_k} satisfying $\Phi(t_{n_k})\phi_{n_k}$ converges to $k \rightarrow \infty$ in \mathbb{X} . Further, we define $\omega(\mathcal{V}) = \{\phi \in \mathbb{X}^+ : \lim_{k \rightarrow \infty} \Phi(t_{n_k})\phi_{n_k} = \phi \text{ for some sequences } \phi_{n_k} \in \mathcal{V}\}$ to be the omega limit set of \mathcal{V} . Based on [33, Lemma 23.1(2)], one learns that $\omega(\mathcal{V})$ is an invariant set, compact, nonempty in \mathbb{X}^+ , and $\omega(\mathcal{V})$ attracts \mathcal{V} . Based on [34, Lemma 2.1(b)], one has

$$\kappa(\Phi(t)\mathcal{V}) \leq \kappa(\omega(\mathcal{V})) + \text{dist}(\Phi(t)\mathcal{V}, \omega(\mathcal{V})) \rightarrow 0, \quad \text{as } t \rightarrow \infty,$$

where $\text{dist}(\Phi(t)\mathcal{V}, \omega(\mathcal{V}))$ represents the distance from $\Phi(t)\mathcal{V}$ to $\omega(\mathcal{V})$. Therefore, $\Phi(t)$ is κ -contracting. It finishes the proof. \square

Combining [35, Theorem 1.1.3(b)], Lemmas 3.3 and 3.4, the below result can be derived.

Theorem 3.1. *The solution semiflow $\Phi(t) : \mathbb{X}^+ \rightarrow \mathbb{X}^+$ of model (2.1) has a global attractor.*

4. Threshold dynamics

4.1. Basic reproduction number

It is now clear that model (2.1) has a disease-free steady state $\mathcal{E}_0 = (S^0(x), V^0(x), 0, 0)$ satisfies

$$\begin{cases} -d_1 \Delta S^0(x) = \Lambda(x) - (\mu(x) + \rho(x))S^0(x) + \theta(x)V^0(x), & x \in \mathbb{D}, \\ -d_2 \Delta V^0(x) = \rho(x)S^0(x) - (\mu(x) + \theta(x))V^0(x), & x \in \mathbb{D}, \\ \frac{\partial S^0(x)}{\partial n} = \frac{\partial V^0(x)}{\partial n} = 0, & x \in \partial \mathbb{D}. \end{cases}$$

The linearized subsystem of (2.1) at \mathcal{E}_0 is

$$\begin{cases} \frac{\partial I}{\partial t} = d_3 \Delta I + \left[\frac{\partial F}{\partial I}(x, S^0(x), 0) + \sigma \frac{\partial F}{\partial I}(x, V^0(x), 0) - (\mu(x) + d(x) + r(x) + \gamma(x)) \right] I \\ \quad + \left[\frac{\partial G}{\partial W}(x, S^0(x), 0) + \sigma \frac{\partial G}{\partial W}(x, V^0(x), 0) \right] W, \quad x \in \mathbb{D}, \quad t > 0, \\ \frac{\partial W}{\partial t} = \alpha(x)I - \xi(x)W, \quad x \in \mathbb{D}, \quad t > 0, \\ \frac{\partial I}{\partial n} = 0, \quad x \in \partial \mathbb{D}, \quad t > 0. \end{cases} \quad (4.1)$$

Under assumption (H₁) and (H₂), the linear system (4.1) is cooperative. Denote $\mathcal{T}(t)$ to be the solution semiflow of (4.1) on $C(\overline{\mathbb{D}}, \mathbb{R}^2)$, where operators are

$$\begin{aligned} \mathcal{A} = & \begin{pmatrix} d_3 \Delta - (\mu(x) + d(x) + r(x) + \gamma(x)) & 0 \\ \alpha(x) & -\xi(x) \end{pmatrix} \\ & + \begin{pmatrix} \frac{\partial F}{\partial I}(x, S^0(x), 0) + \sigma \frac{\partial F}{\partial I}(x, V^0(x), 0) & \frac{\partial G}{\partial W}(x, S^0(x), 0) + \sigma \frac{\partial G}{\partial W}(x, V^0(x), 0) \\ 0 & 0 \end{pmatrix} \\ & := \mathcal{B} + \mathcal{F}. \end{aligned}$$

Allow us to denote $\tilde{\mathcal{T}}$ as the positive semigroup generated by \mathcal{B} . According to [37, Theorem 3.12], one gives the next generator operator

$$\mathcal{L}(\phi)(x) = \int_0^\infty \mathcal{F}(x) \tilde{\mathcal{T}}(t) \phi(x) dt = \mathcal{F}(x) \int_0^\infty \tilde{\mathcal{T}}(t) \phi(x) dt \quad \phi \in C(\overline{\mathbb{D}}, \mathbb{R}^2), \quad x \in \overline{\mathbb{D}}.$$

Define the spectral radius of \mathcal{L} as the basic reproduction number \mathcal{R}_0 , i.e.,

$$\mathcal{R}_0 := r(\mathcal{L}) = \sup\{|\lambda|, \lambda \in \sigma(\mathcal{L})\}.$$

Similar to Refs. [36, Lemma 2.2] and [37], the below consequence is valid.

Lemma 4.1. $\mathcal{R}_0 - 1$ has the identical sign to $s(\mathcal{A})$, where $s(\mathcal{A}) = \sup\{|\lambda|, \lambda \in \sigma(\mathcal{L})\}$ is the spectral bound of \mathcal{A} .

Lemma 4.2. Let $\tilde{\lambda}_0$ satisfy

$$\begin{aligned} d_3 \Delta \phi - (\mu(x) + d(x) + r(x) + \gamma(x))\phi + \tilde{\lambda} \left(\frac{\partial F}{\partial I}(x, S^0(x), 0) \right. \\ \left. + \sigma \frac{\partial F}{\partial I}(x, V^0(x), 0) + \frac{\alpha(x)(\frac{\partial G}{\partial W}(x, S^0(x), 0) + \sigma \frac{\partial G}{\partial W}(x, V^0(x), 0))}{\xi(x)} \right) = 0, \quad x \in \mathbb{D} \end{aligned} \quad (4.2)$$

with $\partial \phi / \partial n = 0$, $x \in \partial \mathbb{D}$, then $\mathcal{R}_0 = 1/\tilde{\lambda}_0$.

Proof. \mathcal{FB}^{-1} is calculated to give

$$\begin{aligned}
& -\mathcal{FB}^{-1}\psi \\
&= -\begin{pmatrix} \frac{\partial F}{\partial I}(x, S^0(x), 0) + \sigma \frac{\partial F}{\partial I}(x, V^0(x), 0) & \frac{\partial G}{\partial W}(x, S^0(x), 0) + \sigma \frac{\partial G}{\partial W}(x, V^0(x), 0) \\ 0 & 0 \end{pmatrix} \\
&\quad \times \begin{pmatrix} (d_3\Delta - (\mu(x) + d(x) + r(x) + \gamma(x)))^{-1} & 0 \\ \alpha(x)(d_3\Delta - (\mu(x) + d(x) + r(x) + \gamma(x)))^{-1}(\xi(x))^{-1} & -(\xi(x))^{-1} \end{pmatrix} \psi \\
&= \begin{pmatrix} -\mathcal{H}(x, S^0(x), V^0(x))(d_3\Delta - (\mu(x) + d(x) + r(x) + \gamma(x)))^{-1} & \mathcal{J}(x, S^0(x), V^0(x)) \\ 0 & 0 \end{pmatrix} \psi,
\end{aligned}$$

where

$$\begin{aligned}
& \mathcal{H}(x, S^0(x), V^0(x)) \\
&= \left[\frac{\partial F}{\partial I}(x, S^0(x), 0) + \sigma \frac{\partial F}{\partial I}(x, V^0(x), 0) + \frac{\alpha(x)}{\xi(x)} \left(\frac{\partial G}{\partial W}(x, S^0(x), 0) + \sigma \frac{\partial G}{\partial W}(x, V^0(x), 0) \right) \right], \\
& \mathcal{J}(x, S^0(x), V^0(x)) = (\xi(x))^{-1} \left[\frac{\partial G}{\partial W}(x, S^0(x), 0) + \sigma \frac{\partial G}{\partial W}(x, V^0(x), 0) \right].
\end{aligned}$$

Due to

$$\begin{aligned}
\mathcal{R}_0 = r(\mathcal{L}) &= r \left(- \left[\frac{\partial F}{\partial I}(x, S^0(x), 0) + \sigma \frac{\partial F}{\partial I}(x, V^0(x), 0) + \frac{\alpha(x)}{\xi(x)} \left(\frac{\partial G}{\partial W}(x, S^0(x), 0) \right. \right. \right. \\
&\quad \left. \left. \left. + \sigma \frac{\partial G}{\partial W}(x, V^0(x), 0) \right) \right] (d_3\Delta - (r(x) + \mu(x) + \gamma(x) + d(x)))^{-1} \right),
\end{aligned}$$

therefore, \mathcal{R}_0 is the principle eigenvalue of

$$\begin{aligned}
& - \left[\frac{\partial F}{\partial I}(x, S^0(x), 0) + \sigma \frac{\partial F}{\partial I}(x, V^0(x), 0) + \alpha(x) \left(\frac{\partial G}{\partial W}(x, S^0(x), 0) + \sigma \frac{\partial G}{\partial W}(x, V^0(x), 0) \right) \right] \\
& \quad \times (d_3\Delta - (r(x) + \mu(x) + \gamma(x) + d(x)))^{-1} (\xi(x))^{-1} \phi = \mathcal{R}_0 \phi, \quad \phi \in C^2(\overline{\mathbb{D}}).
\end{aligned}$$

In other words,

$$\begin{aligned}
& (d_3\Delta \phi - (r(x) + \mu(x) + \gamma(x) + d(x))\phi + \left[\frac{\partial F}{\partial I}(x, S^0(x), 0) + \sigma \frac{\partial F}{\partial I}(x, V^0(x), 0) \right. \\
& \quad \left. + \alpha(x) \left(\frac{\partial G}{\partial W}(x, S^0(x), 0) + \sigma \frac{\partial G}{\partial W}(x, V^0(x), 0) \right) \right] (\xi(x))^{-1} \frac{1}{\mathcal{R}_0} \phi = 0, \quad \phi \in C^2(\overline{\mathbb{D}}).
\end{aligned}$$

It finishes the proof. \square

Remark 4.1. Following Lemma 4.2, \mathcal{R}_0 can be shown by the variational approach of the form

$$\mathcal{R}_0 = \frac{1}{\tilde{\lambda}_0} = \sup_{\phi \in H^1(\mathbb{D}), \phi \neq 0} \left\{ \frac{\int_{\mathbb{D}} \mathcal{H}(x, S^0(x), V^0(x)) \phi^2 dx}{\int_{\mathbb{D}} d_3 |\nabla \phi|^2 + (r(x) + \gamma(x) + d(x) + \mu(x)) \phi^2 dx} \right\}. \quad (4.3)$$

Remark 4.2. Assuming that all parameters of (2.1) are independent of x yields

$$S^0(x) = \frac{\Lambda(\mu + \theta)}{\mu(\mu + \theta + \rho)}, \quad V^0(x) = \frac{\Lambda\rho}{\mu(\mu + \theta + \rho)},$$

In particular, if

$$F(x, S, I) = \frac{\beta_1 SI}{1+qI}, \quad G(x, S, W) = \frac{\beta_2 SW}{1+pW}, \quad F(x, V, I) = \frac{\beta_1 VI}{1+qI}, \quad G(x, V, W) = \frac{\beta_2 VW}{1+pW}, \quad p, q \in [0, 1],$$

then

$$\bar{\mathcal{R}}_0 = \frac{\Lambda(\xi\beta_1 + \alpha\beta_2)(\mu + \theta + \sigma\rho)}{\xi\mu(\mu + \theta + \rho)(\mu + d + r + \gamma)}, \quad (4.4)$$

which will be used in our numerical simulation.

Lemma 4.3. If $\mathcal{R}_0 \geq 1$ ($s(\mathcal{A}) \geq 0$), $s(\mathcal{A})$ is the principal eigenvalue of problem

$$\begin{cases} \lambda\phi_3 = d_3\Delta\phi_3 + \left[-(\mu(x) + d(x) + \gamma(x)) + \sigma\frac{\partial F}{\partial I}(x, V^0(x), 0) + \frac{\partial F}{\partial I}(x, S^0(x), 0) \right]\phi_3 \\ \quad + \left[\sigma\frac{\partial G}{\partial W}(x, V^0(x), 0) + \frac{\partial G}{\partial W}(x, S^0(x), 0) \right]\phi_4, \quad x \in \mathbb{D}, \\ \lambda\phi_4 = -\xi(x)\phi_4 + \alpha(x)\phi_3, \quad x \in \mathbb{D}, \\ \frac{\partial\phi_3}{\partial n} = 0, \quad x \in \partial\mathbb{D}, \end{cases} \quad (4.5)$$

associated with a strongly positive eigenfunction.

Proof. According to (4.1), it can be derived that

$$\begin{cases} I(x, t, \phi) = \Gamma_3(t)\phi_3(t) + \int_0^t \Gamma_3(t-s)\mathcal{P}(I(x, s, \phi), W(x, s, \phi))ds, \\ W(x, t, \phi) = \Gamma_4(t)\phi_4(t) + \int_0^t \Gamma_4(t-s)(\alpha(x)I(x, s, \phi))ds, \end{cases} \quad (4.6)$$

where $\mathcal{P}(x, I, W) = F(x, \phi_1, \phi_3) + G(x, \phi_1, \phi_4) + \sigma F(x, \phi_2, \phi_3) + \sigma G(x, \phi_2, \phi_4) - \gamma(x)\phi_3/(1 + \alpha(x)\phi_3)$.

We rewrite $\tilde{\mathcal{T}}(t)$ as $\tilde{\mathcal{T}}(t) = \tilde{\mathcal{T}}_3(t) + \tilde{\mathcal{T}}_4(t)$, where $\phi = (\phi_3, \phi_4) \in C(\overline{\mathbb{D}}, \mathbb{R}^2)$,

$$\tilde{\mathcal{T}}_3(t)\phi = (0, \Gamma_4(t)\phi_4), \quad \tilde{\mathcal{T}}_4(t)\phi = \left(I(x, t; \phi), \int_0^t \Gamma_4(t-s)(\alpha(x)I(x, s; \phi))ds \right). \quad (4.7)$$

Similar to Ref. [30, Lemma 2.5], $\tilde{\mathcal{T}}_4(t)$ is tight. Therefore, it yields from (4.7) that

$$\sup_{\phi \in C(\overline{\mathbb{D}}, \mathbb{R}^2), \|\phi\| \neq 0} \frac{\|\tilde{\mathcal{T}}_3(t)\phi\|}{\|\phi\|} \leq \sup_{\phi \in C(\overline{\mathbb{D}}, \mathbb{R}^2), \|\phi\| \neq 0} \frac{\|e^{-\pi_4(x)t}\phi_4\|}{\|\phi\|} \leq e^{-\xi^- t}.$$

As a consequence, for every set \mathcal{A} in $C(\overline{\mathbb{D}}, \mathbb{R}^2)$, one has

$$\tau(\tilde{\mathcal{T}}(t)\phi) \leq \tau(\tilde{\mathcal{T}}_3(t)\mathcal{A}) + \tau(\tilde{\mathcal{T}}_4(t)\mathcal{A}) \leq \|\tilde{\mathcal{T}}_3(t)\mathcal{A}\| \leq e^{-\xi^- t}\tau(\mathcal{A}), \quad t > 0.$$

From the above inequality, \mathcal{T} is a τ -contraction on $C(\overline{\mathbb{D}}, \mathbb{R}^2)$ with contraction function $e^{-\xi^- t}$, i.e., the essential spectra radius, $\omega_{ess}(\mathcal{T}(t)) \leq -\xi^-$. Because $\omega_{ess}(\mathcal{T}(t))$ is defined as $\omega_{ess}(\mathcal{T}(t)) := \lim_{t \rightarrow \infty} \vartheta(\mathcal{T}(t))/t$, ϑ is the measure of non-compactness.

As is well known (see Ref. [38]) $\omega = \max\{s(\mathcal{A}), \omega_{ess}(\mathcal{T}(t))\}$, where $\omega := \lim_{t \rightarrow \infty} \ln \|\mathcal{T}(t)\|/t$ is the exponential growth bound of $\mathcal{T}(t)$ such that $\|\mathcal{T}(t)\| \leq M e^{\omega t}$, for $M > 0$. In addition, the spectral radius $r(\mathcal{T}(t))$ of $\mathcal{T}(t)$ fulfills

$$r(\mathcal{T}(t)) = e^{s(\mathcal{A})t} \geq 1, \text{ when } s(\mathcal{A}) \geq 0, \quad t > 0,$$

which means that $\omega(\mathcal{T}(t)) < r(\mathcal{T}(t))$, $t > 0$. Thanks to the generalized Krein-Rutman Theorem (see, Ref. [39, Lemma 2.2]), this concludes the proof. \square

4.2. Stability of steady states

Throughout this subsection, one concentrates on obtaining threshold results for model (2.1) in terms of \mathcal{R}_0 . To begin with, one gives the stability \mathcal{E}_0 for $\mathcal{R}_0 < 1$.

Theorem 4.1. *If $\mathcal{R}_0 < 1$ (or $s(\mathcal{A}) < 0$), the \mathcal{E}_0 is locally asymptotically stable. Further, if $\gamma(x) = 0$, then \mathcal{E}_0 is globally asymptotically stable for $\mathcal{R}_0 < 1$.*

Proof. By analogy with Ref. [36, Theorem 3.1], it is clear that \mathcal{E}_0 is locally asymptotically stable for $\mathcal{R}_0 < 1$, so we just need to prove the global attraction of \mathcal{E}_0 with $\gamma(x) = 0$ in this case. Fix $\epsilon_0 > 0$. From (3.3), there exists $t_1 > 0$ fulfilling that as $t \geq t_1$, $0 \leq S(x, t) \leq S^0(x) + \epsilon_0$, $0 \leq V(x, t) \leq V^0(x) + \epsilon_0$. With the help of the comparison principle yields $(I(x, t), W(x, t)) \leq (\hat{I}(x, t), \hat{W}(x, t))$ on $\overline{\mathbb{D}} \times [t_1, \infty)$, where $(\hat{I}(x, t), \hat{W}(x, t))$ meets

$$\begin{cases} \frac{\partial \hat{I}}{\partial t} = d_3 \Delta \hat{I} + \left(\frac{\partial F}{\partial I}(x, S^0(x) + \epsilon_0, 0) + \sigma \frac{\partial F}{\partial I}(x, V^0(x) + \epsilon_0, 0) - (\mu(x) + d(x)) \right) \hat{I} \\ \quad + \left(\frac{\partial G}{\partial W}(x, S^0(x) + \epsilon_0, 0) + \sigma \frac{\partial G}{\partial W}(x, V^0(x) + \epsilon_0, 0) \right) \hat{W}, \quad x \in \mathbb{D}, \quad t > t_1, \\ \frac{\partial \hat{W}}{\partial t} = \alpha(x) \hat{I} - \xi(x) \hat{W}, \quad x \in \mathbb{D}, \quad t > t_1, \\ \frac{\partial \hat{I}}{\partial n} = \frac{\partial \hat{W}}{\partial n} = 0, \quad x \in \partial \mathbb{D}, \quad t > t_1, \end{cases} \quad (4.8)$$

with $\hat{I}(x, t_1) = I(x, t_1)$, $\hat{W}(x, t_1) = W(x, t_1)$, $x \in \mathbb{D}$. For adequately small $\epsilon_0 > 0$, $s(\mathcal{A}_{\epsilon_0}) < 0$ as $s(\mathcal{A}) < 0$, as well as a corresponding eigenvector $(\psi_3^{\epsilon_0}, \psi_4^{\epsilon_0}) > (0, 0)$. Assume that for $\phi \in \mathbb{X}^+$, one obtains $(I(x, t_1, \phi), W(x, t_1, \phi)) \leq \iota(\psi_3^{\epsilon_0}(x), \psi_4^{\epsilon_0}(x))$ for $x \in \overline{\mathbb{D}}$, $\iota > 0$. Further, we can arrive at

$$(I(x, t_1, \phi), W(x, t_1, \phi)) \leq \iota e^{s(\mathcal{A}_{\epsilon_0}(t-t_1))} (\psi_3^{\epsilon_0}, \psi_4^{\epsilon_0}), \quad t \geq t_1.$$

Thus, $\lim_{t \rightarrow \infty} I(x, t) = 0$, $\lim_{t \rightarrow \infty} W(x, t) = 0$ uniformly for $x \in \overline{\mathbb{D}}$. Furthermore, one can derive $\lim_{t \rightarrow \infty} S(x, t) = S^0(x)$, $\lim_{t \rightarrow \infty} V(x, t) = V^0(x)$ uniformly for $x \in \overline{\mathbb{D}}$. It finishes the proof. \square

Theorem 4.2. *Suppose that $\mathcal{R}_0 > 1$ (or $s(\mathcal{A}) > 0$). There exists $\varepsilon > 0$ satisfying $u_0 = (S_0(x), V_0(x), I_0(x), W_0(x)) \in \mathbb{X}^+$ with $I_0(x) \neq 0$ or $W_0(x) \neq 0$, then $u(x, t; u_0) = (S(x, t), V(x, t), I(x, t), W(x, t))$ satisfies $\liminf_{t \rightarrow \infty} u(x, t; u_0) \geq (\varepsilon, \varepsilon, \varepsilon, \varepsilon)$, uniformly for $x \in \overline{\mathbb{D}}$. Moreover, at least one endemic steady \mathcal{E}^* is included in the model (2.1).*

Proof. Let $\mathbb{X}_0 := \{\phi \in \mathbb{X}^+ : \phi_3(x) \not\equiv 0 \text{ and } \phi_4(x) \not\equiv 0\}$, $\partial\mathbb{X}_0 := \mathbb{X}^+ \setminus \mathbb{X}_0 = \{\phi \in \mathbb{X}^+ : \phi_3(x) \equiv 0 \text{ or } \phi_4(x) \equiv 0\}$, $\mathcal{M}_\partial := \{\phi \in \partial\mathbb{X}_0 : \Phi(t)\phi \in \partial\mathbb{X}_0, t \geq 0\}$, where $\Phi(t) : \mathbb{X}^+ \rightarrow \mathbb{X}^+$ is the semiflow generated by model (2.1). Clearly, $\mathbb{X}^+ = \mathbb{X} \cup \partial\mathbb{X}_0$, where \mathbb{X}_0 is relatively open in \mathbb{X}^+ . Next, we will finish the proof with three claims.

- Claim 1. For $t \geq 0$, $\Phi(t)\mathbb{X}_0 \subseteq \mathbb{X}_0$.

Due to $u_0 = (S_0(x), V_0(x), I_0(x), W_0(x)) \in \mathbb{X}_0$, then $I_0(x) \not\equiv 0$ and $W_0(x) \not\equiv 0$. Let \check{I} fulfill the following equation

$$\frac{\partial \check{I}}{\partial t} = d_3 \Delta \check{I} - \frac{\gamma(x) \check{I}}{1 + a(x) \check{I}} - (r(x) + d(x) + \mu(x)) \check{I}, \quad x \in \mathbb{D}; \quad \frac{\partial \check{I}}{\partial n} = 0, \quad x \in \partial\mathbb{D}, \quad (4.9)$$

with $\check{I}(x, 0) = I(x, 0) = I_0(x)$, $x \in \mathbb{D}$. From the maximum principal and $I_0(x) \not\equiv 0$, one has $\check{I}(x, t) > 0$, for $x \in \overline{\mathbb{D}}, t > 0$. Further, it yields from $\partial I / \partial t \geq d_3 \Delta I - (\mu(x) + d(x))I - \gamma(x)I / (1 + a(x)I)$ and the comparison principal that $I(x, t) \geq \check{I}(x, t) > 0$ $x \in \overline{\mathbb{D}}, t > 0$. Further, by the W -equation of (2.1), one derives

$$W(x, t) = e^{-\xi(x)t} W_0(x) + \int_0^t e^{-\xi(x)(t-s)} \alpha(x) I(x, s) ds. \quad (4.10)$$

This means that for $x \in \overline{\mathbb{D}}, t > 0$, $W(x, t) > 0$. Thus, $\Phi(t)u_0 \in \mathbb{X}_0$, that is, the conclusion in Claim 1 is valid.

- Claim 2. For all $\phi \in \mathcal{M}_\partial$, $\omega(u_0) = \{(S^0(x), V^0(x), 0, 0)\}$, where $\omega(u_0)$ denotes the omega limit set of the orbit $\gamma^+(u_0) := \{\Phi(t)u_0 : t \geq 0\}$.

If $\phi \in \mathcal{M}_\partial$, we have $\Phi(t)\phi \in \partial\mathbb{X}_0$, i.e., $I(x, t) \equiv 0$ or $W(x, t) \equiv 0$. For the former case, based on the W -equation in model (2.1), we still have $\lim_{t \rightarrow \infty} W(x, t) = 0$ uniformly for $x \in \overline{\mathbb{D}}$. Consequently, from the previous two equations of model (2.1), one derives that $\lim_{t \rightarrow \infty} S(x, t) = S^0(x)$, $\lim_{t \rightarrow \infty} V(x, t) = V^0(x)$ uniformly for $x \in \overline{\mathbb{D}}$. For the latter case, $I(x, t^*) \not\equiv 0$ and $W(x, t^*) \equiv 0$ for some $t^* > 0$, utilizing the parabolic maximum principal in the I -equation of model (2.1), then $I(x, t) > 0$ for $x \in \overline{\mathbb{D}}$ and $t > t^*$. However, it is possible to derive $I(x, t) \equiv 0$ from the W -equation of the model (2.1), where $x \in \overline{\mathbb{D}}$ and $t > t^*$; this is a contradiction. Therefore, we can also conclude that $\lim_{t \rightarrow \infty} S(x, t) = S^0(x)$, $\lim_{t \rightarrow \infty} V(x, t) = V^0(x)$ uniformly for $x \in \overline{\mathbb{D}}$. Thus, $\partial\mathcal{M}_0$ is positively invariant relative to $\Phi(x)$.

- Claim 3. $\forall \phi \in \mathbb{X}_0, \limsup_{t \rightarrow \infty} \|\Phi(t)\phi - \mathcal{E}_0\| \geq \delta_0$.

Thanks to the continuity of the principal eigenvalues λ , there is a small enough number $\varepsilon_0 > 0$ meeting $\lambda(\varepsilon_0) + \varepsilon_0 < 0$, where $\lambda(\varepsilon_0)$ satisfies

$$\begin{cases} \lambda\psi = \left[\frac{\gamma(x)}{1 + a(x)\varepsilon_0} - \frac{\partial F}{\partial I} \left(x, S^0(x) - \varepsilon_0, \varepsilon_0 \right) - \sigma \frac{\partial F}{\partial I} \left(x, V^0(x) - \varepsilon_0, \varepsilon_0 \right) + \mu(x) + d(x) + r(x) \right. \\ \quad \left. - \frac{\alpha(x)}{\xi(x)} \left(\frac{\partial G}{\partial W} \left(x, S^0(x) - \varepsilon_0, \varepsilon_0 \right) + \frac{\partial G}{\partial W} \left(x, V^0(x) - \varepsilon_0, \varepsilon_0 \right) \right) \right] \psi - d_3 \Delta \psi, \quad x \in \mathbb{D}, \\ \frac{\partial \psi}{\partial n} = 0, \quad x \in \partial\mathbb{D}, \end{cases} \quad (4.11)$$

and ψ_{ε_0} is the positive eigenvector corresponding to $\lambda(\varepsilon_0)$. Assume that Claim 3 is not valid, then for any $0 < \varepsilon_1 < \varepsilon_0$, $\limsup_{t \rightarrow \infty} \|\Phi(t)\phi - \mathcal{E}_0\| < \varepsilon_1$. So, for $x \in \overline{\mathbb{D}}$ and $t > t_1 > 0$,

$$\begin{aligned} S^0(x) - \varepsilon_0 &< S(x, t) < S^0(x) + \varepsilon_0, \quad 0 < W(x, t) < \varepsilon_0, \\ V^0(x) - \varepsilon_0 &< V(x, t) < V^0(x) + \varepsilon_0, \quad 0 < I(x, t) < \varepsilon_0. \end{aligned} \quad (4.12)$$

Assumptions (H₁) and (H₂) yield

$$\begin{aligned} \frac{F(x, S, I)}{I} &\geq \frac{\partial F}{\partial I}(x, S^0(x) - \varepsilon_0, \varepsilon_0), \quad \frac{F(x, V, I)}{I} \geq \frac{\partial F}{\partial I}(x, V^0(x) - \varepsilon_0, \varepsilon_0), \\ \frac{G(x, S, W)}{W} &\geq \frac{\partial G}{\partial W}(x, S^0(x) - \varepsilon_0, \varepsilon_0), \quad \frac{G(x, V, W)}{W} \geq \frac{\partial G}{\partial W}(x, V^0(x) - \varepsilon_0, \varepsilon_0), \end{aligned}$$

for all $x \in \overline{\mathbb{D}}$. Therefore, for $(S_0(x), V_0(x), I_0(x), W_0(x)) \in \mathbb{X}_0$, there is $\eta > 0$ satisfying $I_0(x) \geq \eta \psi_{\varepsilon_0}(x)$. Combining (4.12) and the arbitrariness of ε_0 , we derive that $I(x, t)$ is the upper solution of the below problem

$$\begin{cases} \frac{\partial \omega}{\partial t} = d_3 \Delta \omega + \left[\frac{\partial F}{\partial I}(x, S^0(x) - \varepsilon_0, \varepsilon_0) + \sigma \frac{\partial F}{\partial I}(x, V^0(x) - \varepsilon_0, \varepsilon_0) \right. \\ \quad \left. + \frac{\alpha(x)}{\xi(x)} \left(\frac{\partial G}{\partial W}(x, S^0(x) - \varepsilon_0, \varepsilon_0) + \sigma \frac{\partial G}{\partial W}(x, V^0(x) - \varepsilon_0, \varepsilon_0) \right) \right. \\ \quad \left. - \left(\frac{\gamma(x)}{1 + a(x)\varepsilon_0} + r(x) + d(x) + \mu(x) \right) \right] \omega, \quad x \in \mathbb{D}, \quad t > t_1, \\ \frac{\partial \psi}{\partial n} = 0, \quad x \in \partial \mathbb{D}, \quad t > t_1; \quad \omega(x, t_1) = \eta \psi_{\varepsilon_0}, \quad x \in \mathbb{D}. \end{cases} \quad (4.13)$$

Evidently, $\eta e^{-\lambda(\varepsilon_0)} \psi_{\varepsilon_0}(x)$ is the only solution to system (4.13). Therefore,

$$I(x, t) \leq \eta e^{-\lambda(\varepsilon_0)} \psi_{\varepsilon_0}(x) \rightarrow \infty \quad \text{uniformly for } x \in \overline{\mathbb{D}}, \text{ as } t \rightarrow \infty.$$

This contradicts Lemma 3.3, which proves the claim.

Comparable to the approach in Ref. [27], define $p(x) : \mathbb{X}^+ \rightarrow [0, \infty)$ for the semiflow $\Phi(t)$ as

$$p(\phi)(x) := \min\{\min_{x \in \overline{\mathbb{D}}} \phi_3(x), \min_{x \in \overline{\mathbb{D}}} \phi_4(x)\}, \quad \forall \phi \in \mathbb{X}^+.$$

Similar to Refs. [27, Theorem 3] and [41, Theorem 3.4], there exists a constant $\delta_1 > 0$ that meets $\min_{\psi \in \omega(\phi)} p(\psi) > \delta_1$, for all $\phi \in \mathbb{X}_0$, which means that $\liminf_{t \rightarrow \infty} I(x, t) \geq \delta_1$, $\liminf_{t \rightarrow \infty} W(x, t) \geq \delta_1$, for $\phi \in \mathbb{X}_0$. Hence, there exists $\delta_2 > 0$ satisfying $\liminf_{t \rightarrow \infty} S(x, t) \geq \delta_2$, $\liminf_{t \rightarrow \infty} V(x, t) \geq \delta_2$ for all $x \in \overline{\mathbb{D}}$. Let $\delta = \min\{\delta_1, \delta_2\}$, then the model is uniformly persistent. By Ref. [34, Remark 3.10 and Theorem 3.7], $\Phi(t) : \mathbb{X}_0 \rightarrow \mathbb{X}_0$ has a global attract \mathcal{E}_0 . According to Ref. [34, Theorem 4.7], model (2.1) has one steady state at least $\mathcal{E}^* = (S^*(x), V^*(x), I^*(x), W^*(x))$. It finishes the proof. \square

5. Bifurcation analysis

Next, using the divergence theory, we will derive a few qualities of positive steady state of model (2.1) by taking the death rate due to disease $d(x) = d$ as a bifurcation parameter. Assuming that

$(S(x), V(x), I(x), W(x))$ is the steady state of model (2.1), then

$$\begin{cases} 0 = d_1 \Delta S + \Lambda(x) - (\mu(x) + \rho(x))S - F(x, S, I) - G\left(x, S, \frac{\alpha(x)}{\xi(x)}I\right) + \theta(x)V, \quad x \in \mathbb{D}, \\ 0 = d_2 \Delta V + \rho(x)S - \sigma F(x, V, I) - \sigma G\left(x, V, \frac{\alpha(x)}{\xi(x)}I\right) - (\mu(x) + \theta(x))V, \quad x \in \mathbb{D}, \\ 0 = d_3 \Delta I + F(x, S, I) + G\left(x, S, \frac{\alpha(x)}{\xi(x)}I\right) + \sigma F(x, V, I) + \sigma G\left(x, V, \frac{\alpha(x)}{\xi(x)}I\right) \\ \quad - (\mu(x) + d + r(x))I - \frac{\gamma(x)I}{1 + a(x)I}, \quad x \in \mathbb{D}, \\ 0 = \frac{\partial S}{\partial n} = \frac{\partial V}{\partial n} = \frac{\partial I}{\partial n}, \quad x \in \partial\mathbb{D}, \end{cases} \quad (5.1)$$

and $W(x) = \alpha(x)I/\xi(x)$. Obviously, $(S^0(x), V^0(x), 0)$ fulfills the Eq (5.1). Denote d^* to be the principal eigenvalue of the below equation

$$\begin{cases} d\psi = d_3 \Delta \psi + \left[\frac{\partial F}{\partial I}(x, S^0(x), 0) + \frac{\alpha(x)}{\xi(x)} \frac{\partial G}{\partial I}(x, S^0(x), 0) + \sigma \frac{\partial F}{\partial I}(x, V^0(x), 0) \right. \\ \quad \left. + \sigma \frac{\alpha(x)}{\xi(x)} \frac{\partial G}{\partial I}(x, V^0(x), 0) - (\mu(x) + r(x) + \gamma(x)) \right] \psi, \quad x \in \mathbb{D}, \\ 0 = \frac{\partial \psi}{\partial n}, \quad x \in \partial\mathbb{D}, \end{cases} \quad (5.2)$$

and the corresponding positive eigenfunction $\psi_0(x)$ meeting $\max_{x \in \mathbb{D}} \psi_0(x) = 1$. Moreover, it also realizes that $d = d^*$ is equivalent to $\mathcal{R}_0 = 1$ or $\tilde{\lambda} = 1$. Let

$$\begin{aligned} \mathcal{L}(x) = & \frac{\partial F}{\partial I}(x, S^0(x), 0) + \frac{\alpha(x)}{\xi(x)} \frac{\partial G}{\partial I}(x, S^0(x), 0) + \sigma \frac{\partial F}{\partial I}(x, V^0(x), 0) \\ & + \sigma \frac{\alpha(x)}{\xi(x)} \frac{\partial G}{\partial I}(x, V^0(x), 0) - (\mu(x) + r(x) + \gamma(x)). \end{aligned} \quad (5.3)$$

If $\mathcal{L}(x) \equiv \mathcal{L}$ is a constant, then $b^* = \mathcal{L}$. In the following, we investigate the scenario where $\mathcal{L}(x) \neq$ is a constant and it may vary in sign in \mathbb{D} . Analyze the below problem

$$\begin{cases} \Delta \tilde{\varphi}(x) + \Lambda \mathcal{L}(x) \tilde{\varphi}(x) = 0, \quad x \in \mathbb{D}, \\ \frac{\partial \tilde{\varphi}}{\partial n} = 0, \quad x \in \partial\mathbb{D}. \end{cases} \quad (5.4)$$

By Ref. [42, Theorem 4.2], (5.4) admits a nonzero principal eigenvalue $\Lambda_0 = \Lambda(\mathcal{L})$ if and only if \mathcal{L} can change the sign and $\int_{\mathbb{D}} \mathcal{L}(x) dx \neq 0$.

Regarding the sign problem of the principal eigenvalue d^* , our results are as follows.

Lemma 5.1. *The principal eigenvalue d^* of (5.4) satisfies the following characteristics*

- (i) if $\int_{\mathbb{D}} \mathcal{L}(x) dx \geq 0$, then $d^* > 0$ for all $d_3 > 0$;
- (ii) if $\int_{\mathbb{D}} \mathcal{L}(x) dx < 0$, then $d^* > 0$ for $d_3 < 1/\Lambda(\mathcal{L})$; $d^* < 0$ for $d_3 > 1/\Lambda(\mathcal{L})$.

Now, we process considering d as the bifurcation parameter and studying the local branch of the positive solution of (5.1), which branches from the branch of $\{(S^0(x), V^0(x), 0, d) : d \geq 0\}$. At first, from the transformation $u = S$, $\omega = V$, $v = I$, Eq (5.1) can rewritten as

$$\begin{cases} 0 = d_1 \Delta u + \Lambda(x) - (\mu(x) + \rho(x))u - F(x, u, v) - G(x, u, \frac{\alpha(x)}{\xi(x)}v) + \theta(x)\omega, & x \in \mathbb{D}, \\ 0 = d_2 \Delta \omega + \rho(x)u - \sigma F(x, \omega, v) - \sigma G\left(x, \omega, \frac{\alpha(x)}{\xi(x)}v\right) - (\mu(x) + \theta(x))\omega, & x \in \mathbb{D}, \\ 0 = d_3 \Delta v + F(x, u, v) + G\left(x, u, \frac{\alpha(x)}{\xi(x)}v\right) + \sigma F(x, \omega, v) + \sigma G\left(x, \omega, \frac{\alpha(x)}{\xi(x)}v\right) \\ \quad - (\mu(x) + d + r(x))v - \frac{\gamma(x)v}{1 + a(x)v}, & x \in \mathbb{D}, \\ 0 = \frac{\partial u}{\partial n} = \frac{\partial \omega}{\partial n} = \frac{\partial v}{\partial n}, & x \in \partial \mathbb{D}. \end{cases} \quad (5.5)$$

For $p > n$, let $\mathcal{X} = \{u, \omega \in W^{2,p}(\mathbb{D}) : \partial u(x)/\partial n = \partial \omega(x)/\partial n = 0\}$ and $\mathcal{Y} = L^p(\mathbb{D})$. Define

$$\mathbb{B} = \{(u, \omega, v, d) \in \mathcal{X} \times \mathcal{X} \times \mathcal{X} \times \mathbb{R}_+ : (u, \omega, d) \text{ is a positive solution of (5.1)}\}.$$

Theorem 5.1. *Let d^* be the principal eigenvalue of problem (5.1).*

(i) *There is a connected component \mathbb{B}_1 of $\overline{\mathbb{B}}$ including $(u, \omega, 0, d^*)$, and the projection $\text{proj}_d \mathbb{B}_1$ of \mathbb{B}_1 into the d -axis meets $(0, d^*] \subset \text{proj}_d \mathbb{B}_1 \subset (0, C]$ for*

$$C = \max_{x \in \mathbb{D}} \left\{ \frac{\partial F}{\partial I}(x, S^0(x), 0) + \frac{\alpha(x)}{\xi(x)} \frac{\partial G}{\partial I}(x, S^0(x), 0) \right. \\ \left. + \sigma \frac{\partial F}{\partial I}(x, V^0(x), 0) + \sigma \frac{\alpha(x)}{\xi(x)} \frac{\partial G}{\partial I}(x, V^0(x), 0) \right\}. \quad (5.6)$$

Specifically, for $0 < d < d^$, Eq (5.1) has a positive steady state solution at least.*

(ii) *Near $d = d^*$, \mathbb{B}_1 is a smooth curve $E_1 = \{(u(s), \omega(s), v(s), d(s)) : s \in (0, \varepsilon)\}$, where $u(s) = u_1^* + s\phi_0(s) + o(s)$, $\omega(s) = \omega_1^* + s\chi_0(s) + o(s)$, and $v(s) = s\psi_0(s) + o(s)$. Here, $\psi_0(x) > 0$ is the principal eigenvalue and satisfies (5.2), and $(\phi_0(x), \chi_0(x)) < (0, 0)$ fulfills*

$$\begin{cases} 0 = d_1 \Delta \phi_0(x) - (\mu(x) + \rho(x))\phi_0(x) + \theta(x)\chi_0(x) \\ \quad - \left[\frac{\partial F}{\partial I}(x, S^0(x), 0) + \frac{\alpha(x)}{\xi(x)} \frac{\partial G}{\partial I}(x, S^0(x), 0) \right] \psi_0(x), & x \in \mathbb{D}, \\ 0 = d_2 \Delta \chi_0(x) - (\mu(x) + \theta(x))\chi_0(x) + \rho(x)\phi_0(x) \\ \quad - \sigma \left[\frac{\partial F}{\partial I}(x, V^0(x), 0) + \frac{\alpha(x)}{\xi(x)} \frac{\partial G}{\partial I}(x, V^0(x), 0) \right] \psi_0(x), & x \in \mathbb{D}, \\ 0 = \frac{\partial \phi_0(x)}{\partial n} = \frac{\partial \chi_0(x)}{\partial n}, & x \in \partial \mathbb{D}. \end{cases} \quad (5.7)$$

Further, $d'(0) = \mathcal{N}/(\int_{\mathbb{D}} \psi_0^2(x)dx)$, where ' stands for derivative and

$$\begin{aligned} \mathcal{N} = & \left[\frac{\partial^2 F}{\partial v^2}(x, S^0, 0) + \left(\frac{\alpha(x)}{\xi(x)} \right)^2 \frac{\partial^2 G}{\partial v^2}(x, S^0, 0) \right] \psi_0^3 + 2 \left[\frac{\partial^2 F}{\partial u \partial v}(x, S^0, 0) \right. \\ & \left. + \frac{\alpha(x)}{\xi(x)} \frac{\partial^2 G}{\partial u \partial v}(x, S^0, 0) \right] \phi_0 \psi_0^2 + \sigma \left[\frac{\partial^2 F}{\partial v^2}(x, V^0, 0) + \left(\frac{\alpha(x)}{\xi(x)} \right)^2 \frac{\partial^2 G}{\partial v^2}(x, V^0, 0) \right] \psi_0^3 \\ & + 2\sigma \left[\frac{\partial^2 F}{\partial u \partial v}(x, V^0, 0) + 2 \frac{\alpha(x)}{\xi(x)} \frac{\partial^2 G}{\partial u \partial v}(x, V^0, 0) \right] \phi_0 \psi_0^2. \end{aligned} \quad (5.8)$$

Proof. Similar to the approach in Ref. [43], denote $\mathcal{G} : \mathcal{X} \times \mathcal{X} \times \mathcal{X} \times \mathbb{R} \rightarrow \mathcal{Y} \times \mathcal{Y} \times \mathcal{Y}$ by

$$\mathcal{G}(u, \omega, v) = \begin{pmatrix} d_1 \Delta u + \Lambda(x) - (\mu(x) + \rho(x))u - F(x, u, v) - G(x, u, \frac{\alpha(x)}{\xi(x)}v) + \theta(x)\omega \\ d_2 \Delta v + \rho(x)u - \sigma F(x, u, v) - \sigma G(x, \omega, \frac{\alpha(x)}{\xi(x)}v) - (\mu(x) + \theta(x))\omega \\ d_3 \Delta + F(x, u, v) + G(x, u, \frac{\alpha(x)}{\xi(x)}v) + \sigma F(x, \omega, v) \\ \quad + \sigma G(x, \omega, \frac{\alpha(x)}{\xi(x)}v) - (\mu(x) + d)v - \frac{\gamma(x)v}{1+a(x)v} \end{pmatrix}.$$

Taking partial derivative with respect to (u, ω, v) , we can get

$$\begin{aligned} & \mathcal{G}_{(u, \omega, v)}(S^0, V^0, 0, \gamma^*)[\phi, \chi, \psi] \\ = & \begin{pmatrix} d_1 \Delta \phi - (\mu(x) + \rho(x))\phi - \left(\frac{\partial F}{\partial v}(x, S^0, 0) + \frac{\alpha(x)}{\xi(x)} \frac{\partial G}{\partial v}(x, S^0, 0) \right) \psi + \theta(x)\chi \\ d_2 \Delta \chi + \rho(x)\phi - \sigma \left(\frac{\partial F}{\partial v}(x, V^0, 0) + \frac{\alpha(x)}{\xi(x)} \frac{\partial G}{\partial v}(x, V^0, 0) \right) \psi - (\mu(x) + \theta(x))\chi \\ d_3 \Delta \psi + \mathcal{L}(x)\psi - b^* \psi \end{pmatrix}. \end{aligned} \quad (5.9)$$

Moreover, calculating the second-order partial derivatives for \mathcal{G} about (u, ω, v) leads to

$$\begin{aligned} & \mathcal{G}_{(u, \omega, v), (u, \omega, v)}(S^0, V^0, 0, d^*)[\phi, \chi, \psi]^2 \\ = & \begin{pmatrix} - \left[\frac{\partial^2 F}{\partial v^2}(x, S^0, 0) + \left(\frac{\alpha(x)}{\xi(x)} \right)^2 \frac{\partial^2 G}{\partial v^2}(x, S^0, 0) \right] \psi^2 - 2 \left[\frac{\partial^2 F}{\partial u \partial v}(x, S^0, 0) + \frac{\alpha(x)}{\xi(x)} \frac{\partial^2 G}{\partial u \partial v}(x, S^0, 0) \right] \phi \psi \\ - \sigma \left[\frac{\partial^2 F}{\partial v^2}(x, V^0, 0) + \left(\frac{\alpha(x)}{\xi(x)} \right)^2 \frac{\partial^2 G}{\partial v^2}(x, V^0, 0) \right] \psi^2 - 2\sigma \left[\frac{\partial^2 F}{\partial u \partial v}(x, V^0, 0) + 2 \frac{\alpha(x)}{\xi(x)} \frac{\partial^2 G}{\partial u \partial v}(x, V^0, 0) \right] \phi \psi \\ \left[\frac{\partial^2 F}{\partial v^2}(x, S^0, 0) + \left(\frac{\alpha(x)}{\xi(x)} \right)^2 \frac{\partial^2 G}{\partial v^2}(x, S^0, 0) - \gamma(x)a(x) \right] \psi^2 + 2 \left[\frac{\partial^2 F}{\partial u \partial v}(x, S^0, 0) + \frac{\alpha(x)}{\xi(x)} \frac{\partial^2 G}{\partial u \partial v}(x, S^0, 0) \right] \phi \psi \\ + \sigma \left[\frac{\partial^2 F}{\partial v^2}(x, V^0, 0) + \left(\frac{\alpha(x)}{\xi(x)} \right)^2 \frac{\partial^2 G}{\partial v^2}(x, V^0, 0) \right] \psi^2 + 2\sigma \left[\frac{\partial^2 F}{\partial u \partial v}(x, V^0, 0) + 2 \frac{\alpha(x)}{\xi(x)} \frac{\partial^2 G}{\partial u \partial v}(x, V^0, 0) \right] \phi \psi \end{pmatrix}. \end{aligned}$$

Therefore, it's convenient to check that the core $\mathcal{G}(u, \omega, v)(S^0, V^0, 0, b^*) = \text{span}\{\psi_0, \chi_0, \phi_0\}$, with ϕ_0 as the positive eigenfunction of (5.2), (ϕ_0, χ_0) fulfills (5.7). Based on the Lemma 3.2, $(S^0(x), V^0(x))$ is globally asymptotically stable in $C(\overline{\mathbb{D}}, \mathbb{R})$. This indicates that inverse $[d_2 \Delta - (\mu(x) + \theta(x))]^{-1}$ and $[d_1 \Delta - (\mu(x) + \rho(x))]^{-1}$ exist and are positive operators. Hence, $\phi_0(x) < 0$ and $\chi_0(x) < 0$ for $x \in \mathbb{D}$.

We next consider the range

$$\text{range } \mathcal{G}_{(u, \omega, v)}(S^0, V^0, 0, b^*) = \left\{ (z_1, z_2, z_3 \in \mathcal{Y}^3) : \int_{\mathbb{D}} z_3(x) \psi_0(x) dx = 0 \right\}. \quad (5.10)$$

It is convenient to observe that $(z_1, z_2, z_3) \in \text{range } \mathcal{G}_{(u, \omega, v)}(S^0, V^0, 0, b^*)$ if and only if there has $(\phi, \chi, \psi) \in \mathcal{X} \times \mathcal{X} \times \mathcal{X}$ satisfying

$$\begin{aligned}
z_1 &= d_1 \Delta \phi - (\mu(x) + \rho(x))\phi - \left(\frac{\partial F}{\partial \nu}(x, S^0, 0) + \frac{\alpha(x)}{\xi(x)} \frac{\partial G}{\partial \nu}(x, S^0, 0) \right) \psi + \theta(x)\chi, \\
z_2 &= d_2 \Delta \chi + \rho(x)\phi - \sigma \left(\frac{\partial F}{\partial \nu}(x, V^0, 0) + \frac{\alpha(x)}{\xi(x)} \frac{\partial G}{\partial \nu}(x, V^0, 0) \right) \psi - (\mu(x) + \theta(x))\chi, \\
z_3 &= d_3 \Delta \psi + \mathcal{L}(x)\psi - b^* \psi.
\end{aligned}$$

Hence,

$$\int_{\mathbb{D}} z_3(x) \psi_0(x) dx = d_3 \int_{\mathbb{D}} \Delta \psi(x) \psi_0(x) dx + \int_{\mathbb{D}} (\mathcal{L}(x)\psi - b^* \psi) \psi(x) dx. \quad (5.11)$$

Combining the integration by parts and the boundary conditions, we can derive $\int_{\mathbb{D}} \Delta \psi(x) \psi_0(x) dx = \int_{\mathbb{D}} \Delta \psi_0(x) \psi(x) dx$. Further, from (5.2) and (5.11), we can obtain $\int_{\mathbb{D}} z_3(x) \psi_0(x) dx = 0$, which in contrast implicates that (5.10) is valid. Since

$$\mathcal{G}_{(u,\omega,\nu),b}(S^0, V^0, 0, b^*)[\phi_0, \chi_0, \psi_0] = (0, 0, -\psi_0),$$

and $\int_{\mathbb{D}} [-\psi_0(x)] \psi_0(x) dx < 0$, then $\mathcal{G}_{(u,\omega,\nu),b}(S^0, V^0, 0, b^*)[\phi_0, \chi_0, \psi_0] \notin \text{range} \mathcal{G}_{(u,\omega,\nu)}(S^0, V^0, 0, b^*)$. Using the bifurcation theorem for simple eigenvalues from Ref. [44], it is derived that the positive solution set of (5.7) in $(S^0, V^0, 0, b^*)$ is a curve of E_1 , where $(u'(0), \omega'(0), \nu'(0)) = (\phi_0, \chi_0, \psi_0)$. According to Ref. [45], we launch $b'(0)$ in the following form

$$b'(0) = -\frac{\langle l, \mathcal{G}_{(u,\omega,\nu),(u,\omega,\nu)}(S^0, V^0, 0, b^*)[\phi_0, \chi_0, \psi_0] \rangle^2}{2 \langle l, \mathcal{G}_{(u,\omega,\nu),b}(S^0, V^0, 0, b^*)[\phi_0, \chi_0, \psi_0] \rangle},$$

where l is defined as $\langle l, [z_1, z_2, z_3] \rangle = \int_{\mathbb{D}} z_3 \psi_0(x) dx$. By direct computing, one can conclude that the second component of $\mathcal{G}_{(u,\omega,\nu),(u,\omega,\nu)}(S^0, V^0, 0, d^*)[\phi_0, \chi_0, \psi_0]^2$ takes the form

$$\begin{aligned}
\mathcal{G}(x) &= \left[\frac{\partial^2 F}{\partial \nu^2}(x, S^0, 0) + \left(\frac{\alpha(x)}{\xi(x)} \right)^2 \frac{\partial^2 G}{\partial \nu^2}(x, S^0, 0) - \gamma(x)a(x) \right] \psi_0^2 + 2 \left[\frac{\partial^2 F}{\partial u \partial \nu}(x, S^0, 0) \right. \\
&\quad \left. + \frac{\alpha(x)}{\xi(x)} \frac{\partial^2 G}{\partial u \partial \nu}(x, S^0, 0) \right] \phi_0 \psi_0 + \sigma \left[\frac{\partial^2 F}{\partial \nu^2}(x, V^0, 0) + \left(\frac{\alpha(x)}{\xi(x)} \right)^2 \frac{\partial^2 G}{\partial \nu^2}(x, V^0, 0) \right] \psi_0^2 \\
&\quad + 2\sigma \left[\frac{\partial^2 F}{\partial u \partial \nu}(x, V^0, 0) + 2 \frac{\alpha(x)}{\xi(x)} \frac{\partial^2 G}{\partial u \partial \nu}(x, V^0, 0) \right] \phi_0 \psi_0.
\end{aligned}$$

Hence,

$$b'(0) = \frac{\int_{\mathbb{D}} \mathcal{G}_0(x) \psi_0(x) dx}{2 \int_{\mathbb{D}} \psi_0^2(x) dx} = \frac{\mathcal{N}}{\int_{\mathbb{D}} \psi_0^2(x) dx},$$

where \mathcal{N} is defined as in (5.8). Similar to the methods in Ref.[19, Theorem 3.1] and [18, Theorem 5.3], it's verified that all conditions of Ref.[43, Theorem 4.4] are fulfilled. Therefore, the branching is generated around $(S^0, V^0, 0)$ as $\mathcal{R}_0 = 1$. \square

6. Numerical simulations

Throughout this subsection, one conducts fits to account for the impacts of spatially heterogeneous parameters and individuals diffusion on disease propagation. In the interest of simplicity, we take the domain to be $\mathbb{D} = [0, 20]$, and set $d_1 = 0.02$, $d_2 = 0.05$, and $d_3 = 0.005$ to reflect that the individual's mobility is impacted as a result of the disease. Specifically, we consider the general incidence functions $F(x, S, I) = \beta_1(x)SI/(1 + qI)$, $G(x, S, W) = \beta_2(x)SW/(1 + pW)$, $F(x, V, I) = \beta_1(x)VI/(1 + qI)$, $G(x, V, W) = \beta_2(x)V/(1 + pW)$, where $\beta_i(x) \in C^2(\bar{\mathbb{D}})$. According to [26, 48], let's select the parameters $\mu(x) = 4.7 \times 10^{-5} + 2.35 \times 10^{-5} \sin 2x$, $d(x) = 3 \times 10^{-4} + 3 \times 10^{-5} \sin 2x$, $\alpha(x) = 50 + 50 \sin 2x$, $\xi(x) = 0.02 + 0.02 \sin 2x$, $r(x) = 0.25 + 0.2 \sin 2x$. The other parameters will be selected depending on the model.

To begin, we select $\sigma = 0.01$, $q = 2 \times 10^{-6}$, $p = 1 \times 10^{-6}$, $\Lambda(x) = 15 + 7.5 \sin 2x$, $\rho(x) = 4 \times 10^{-3} + 2 \times 10^{-3} \sin 2x$, $\theta(x) = 1.4 \times 10^{-4} + 7 \times 10^{-5} \sin 2x$, $a(x) = 1.75 \times 10^{-3} + 8.75 \times 10^{-4} \sin 2x$, $\beta_1(x) = 1.5 \times 10^{-9} + 7.5 \times 10^{-10} \sin 2x$, $\beta_2(x) = 1.88 \times 10^{-9} + 9.4 \times 10^{-10} \sin 2x$, $\gamma(x) = 0$. Other parameters are shown above, and we select

$$\mathcal{U}(x) = \begin{pmatrix} 86460 - 400 \cos 2x \\ 230000 - 800 \cos 2x \\ 5 - 0.5 \cos 2x \\ 200 - 20 \cos 2x \end{pmatrix}, \quad \forall x \in [0, 20], \quad \mathcal{U} = (S_0, V_0, I_0, B_0)^T.$$

We apply the numerical method mentioned in Ref.[36] to calculate $\mathcal{R}_0 \approx 0.9901 < 1$, which means the disease will ultimately become extinct. As a matter of fact, one can verify in Figure 2(a) and (b) that as time t evolves, $I(x, t)$ and $W(x, t)$ tends to zero, which is compatible with the result that Theorem 4.1.

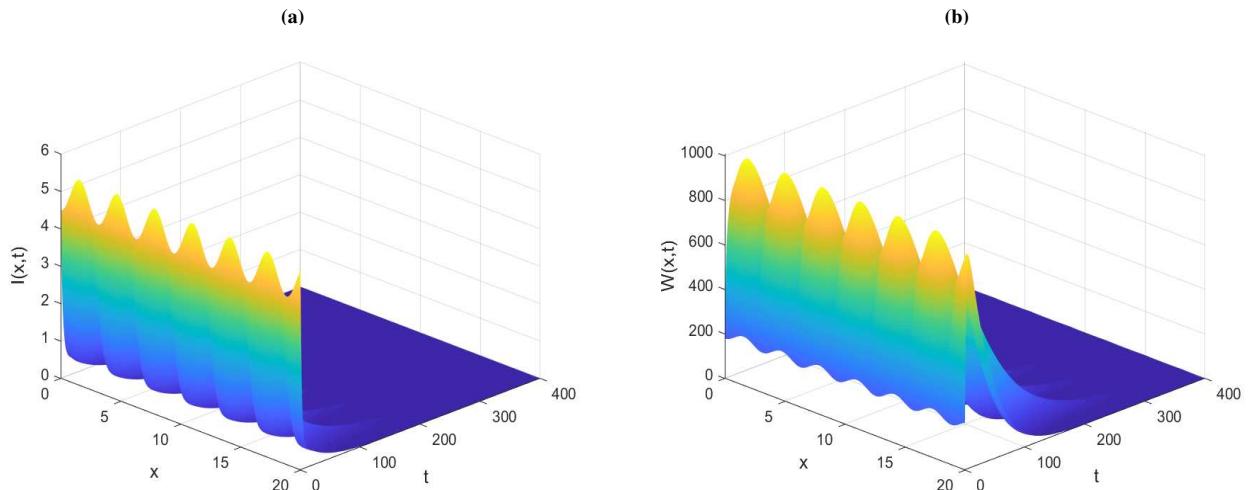


Figure 2. The spatio-temporal distribution of $I(x, t)$ and $W(x, t)$ with $\mathcal{R}_0 \approx 0.9901$: (a) $I(x, t)$; (b) $W(x, t)$.

If we alter the parameters $r(x) = 0.012 + 0.0096 \sin 2x$, $\gamma(x) = 0.03 + 0.0015 \sin 2x$, and $\xi(x) = 0.02 + 0.01 \sin 2x$, the rest of the values are the same as in Figure 2. In this scenario, we derive $\mathcal{R}_0 \approx 2.4875 > 1$. It follows that Theorem 4.2 shows the illness is persistently present. This is shown in Figure 3(a) and (b),

where $I(x, t)$, $W(x, t)$ are periodic oscillations in the whole region. From Figure 3(c) and (d), it can also be found that because of the spatial heterogeneity, $I(x, t)$ and $W(x, t)$ vary geographically across time.

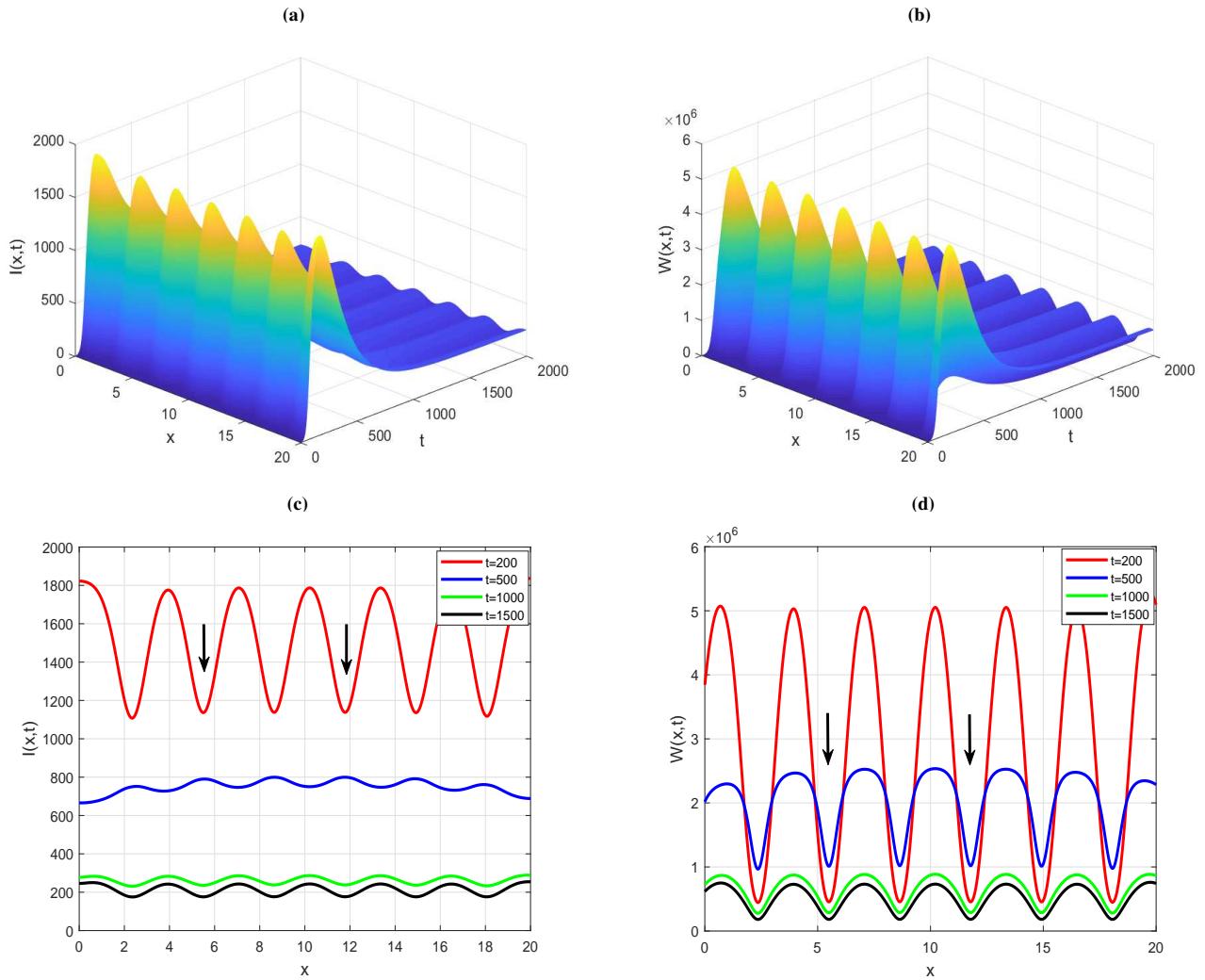


Figure 3. The effect on disease propagation in the case where $\mathcal{R}_0 \approx 2.4875 > 1$: (a)–(b): spatio-temporal evolution of $I(x, t)$ and $W(x, t)$; (c)–(d): regional differences in the distribution of $I(x, t)$ and $W(x, t)$ under different times.

Next, we turn to the influence of spatially heterogeneous parameters on disease propagation. In Figure 4(a)–(c), one illustrates how the vaccination rate $\rho(x)$ affects the quantity of $S(x, t)$, $V(x, t)$, and $I(x, t)$ as time $t = 1500$. In this case, let's choose $\rho(x) = 4 \times 10^{-3} + 4 \times 10^{-3}\rho_1 \sin 2x$, with ρ_1 gradually increasing from 0, 0.2, 0.3, to 0.5. With a growing heterogeneity in vaccination rates, $S(x, t)$ and $V(x, t)$ show large regional diversity. The number of infections in areas with high vaccination rates is known to be relatively low due to the high number of vaccinations. Nevertheless, Figure 4(c) also shows in the same region, e.g., $x \in [12, 14]$, the amount of infected individuals doesn't oscillate significantly as ρ_1 increases, possibly due to the fact that vaccinated individuals remain at high risk of infection. Consequently, along with large-scale immunization, we should also concentrate on the

effectiveness of the vaccine. From Figure 4(d)–(f), it is not uncommon to notice that similar results can be obtained when the model parameters $\theta(x)$ spatial heterogeneity are enhanced. Additionally, from Figure 4(g)–(i), one can observe that when the spatial heterogeneity of the maximum treatment rate $\gamma(x) = 0.03 + 0.03\gamma_1 \sin 2x$ changes, i.e., γ_1 increases from 0, 0.25, 0.5, to 0.75, the regional variability of the regional variability of $S(x, 1500)$, $I(x, 1500)$, and $W(x, 1500)$ will be smaller. In Figure 4(h), one could notice that at the identical location, e.g., $x \in [6, 8]$, the peak of infected individuals decreases as γ_1 increases, which means that by changing the heterogeneous intensity of the maximum treatment rate, the peak of the disease outbreak can be reduced to some extent. Simultaneously, in Figure 4(j)–(i), it can be noticed that when the spatial heterogeneity intensity ξ_1 of $\xi(x) = 0.02 + 0.02\xi_1 \sin 2x$ gradually increases from 0, 0.2, 0.4, to 0.6, we can also derive similar results as in Figure 4(g)–(i). This also reinforces the fact that improving local water sanitation and personal hygiene practices are also vitally important for disease control.

Further, let's clarify the way in which the diffusion coefficient influences the propagation of the illness. In Figure 5(a) and (b), variation of distribution of $S(x, 1500)$ for diffusion rates of $d_1 = 0.2$ and 0.002 is shown. This means that as d_1 increases, the $S(x, t)$ gets more uniform throughout the region. Figure 5(c) and (d) also display the two scenarios for $S(x, t)$ at $t = 1500$ with diffusion rates $d_3 = 0.5$ and $d_3 = 0.005$. By comparing Figure 5(c) and (d), it is apparent that as the diffusion coefficient d_3 increases, the infected individuals present a homogeneous distribution throughout the field. Mathematical simulation outcomes indicate that the propagation of individuals can alter the local spatial distribution of the illness to some extent, and limiting the cross-regional movement of infected individuals during an epidemic is among the least powerful methods of controlling the illness.

In addition, we pay attention to how spatial heterogeneity contributes to the basic reproduction number \mathcal{R}_0 . Here, let's choose the parameters $\xi(x) = 0.048 + 0.048c \sin kx$, $\beta_1(x) = 1.5 \times 10^{-9} + 1.5 \times 10^{-9}c \sin kx$, $\beta_2(x) = 1.88 \times 10^{-9} + 1.88 \times 10^{-9} \sin kx$, where $0 \leq c \leq 1$ and $k = 2, 4, 6$. Other parameters are the same as in the Figure 3. As illustrated by the graphs in Figure 6(a)–(c), variations in the spatial heterogeneity parameters $\beta_1(x)$ and $\beta_2(x)$ increase or decrease the risk of illness propagation, and by comparing Figure 6(b) and (c), it can be observed that \mathcal{R}_0 has different monotonicity for c as k takes different values. The above simulation results also indicated that overlooking spatial heterogeneity may result in misclassification of illness propagation.

Lastly, let's look at the link between \mathcal{R}_0 and the main parameters in the model. Here, we pick the parameters $\Lambda(x) = \Lambda + 1.5 \sin 2x$, $\rho(x) = \rho + 2 \times 10^{-3} \sin 2x$, $\theta(x) = \theta + 7 \times 10^{-5} \sin 2x$, $\mu(x) = \mu + 2.25 \times 10^{-5} \sin 2x$, $\beta_1(x) = \beta_1 + 1.5 \times 10^{-6} \sin 2x$, $d(x) = d + 0.08 \sin 2x$, $\beta_2(x) = \beta_2 + 3.135 \times 10^{-6} \sin 2x$, $r(x) = r + 0.0096 \sin 2x$, $\xi(x) = \xi + 0.01 \sin 2x$, $\gamma(x) = \gamma + 0.0015 \sin 2x$, $\alpha(x) = \alpha + 50 \sin 2x$. Based on the methods in [49], we choose $\Lambda = 15$, $\rho = 4 \times 10^{-3}$, $\theta = 1.4 \times 10^{-4}$, $\mu = 4.5 \times 10^{-5}$, $d = 0.1$, $\beta_1 = 3 \times 10^{-6}$, $\beta_2 = 6.27 \times 10^{-6}$, $r = 0.012$, $\gamma = 0.03$, $\xi = 0.02$, $\alpha = 50$, and the sensitivity indices for each parameter can be calculated separately for \mathcal{R}_0 . As shown in Figure 7, \mathcal{R}_0 has the largest sensitivity index in relation to $\xi(x)$ and $\Lambda(x)$, followed by $\beta_2(x)$, $\alpha(x)$, $\rho(x)$, $\mu(x)$, $d(x)$, $\theta(x)$, $\gamma(x)$, $r(x)$, $\beta_1(x)$. We also observe that \mathcal{R}_0 is positively associated with the variables $\theta(x)$, $\beta_1(x)$, $\beta_2(x)$, $\alpha(x)$, $\Lambda(x)$, and those with $r(x)$, $d(x)$, $\gamma(x)$, $\rho(x)$, $\xi(x)$, $\mu(x)$ are negatively correlated. The above results reveal that it is necessary to disinfect contaminated environments in outbreak areas in a timely manner, and to seal off and control areas with frequent outbreaks to reduce the movement of people. At the same time, we should raise the awareness of the local people on self-prevention, such as drinking healthy and hygienic drinking water and maintaining good hygienic habits.

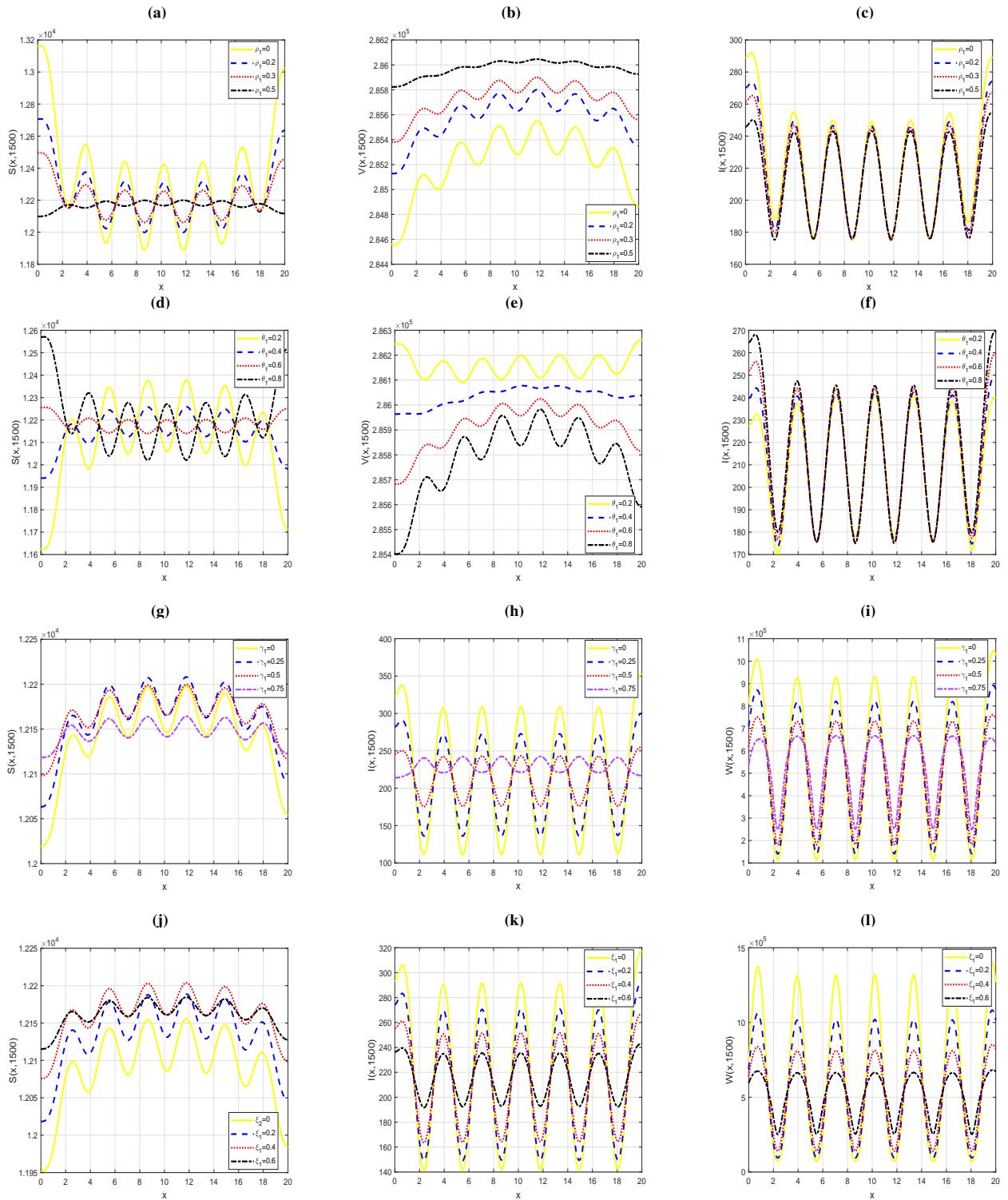


Figure 4. Influence of spatially heterogeneous parameters on the disease distribution of model (2.1), (a)-(c) $\rho(x)$ on $S(x, 1500)$, $V(x, 1500)$, $I(x, 1500)$; (d)-(f) $\theta(x)$ on $S(x, 1500)$, $V(x, 1500)$, $I(x, 1500)$; (g)-(i) $\gamma(x)$ on $S(x, 1500)$, $I(x, 1500)$, $W(x, 1500)$; (j)-(l) $\xi(x)$ on $S(x, 1500)$, $I(x, 1500)$, $W(x, 1500)$.

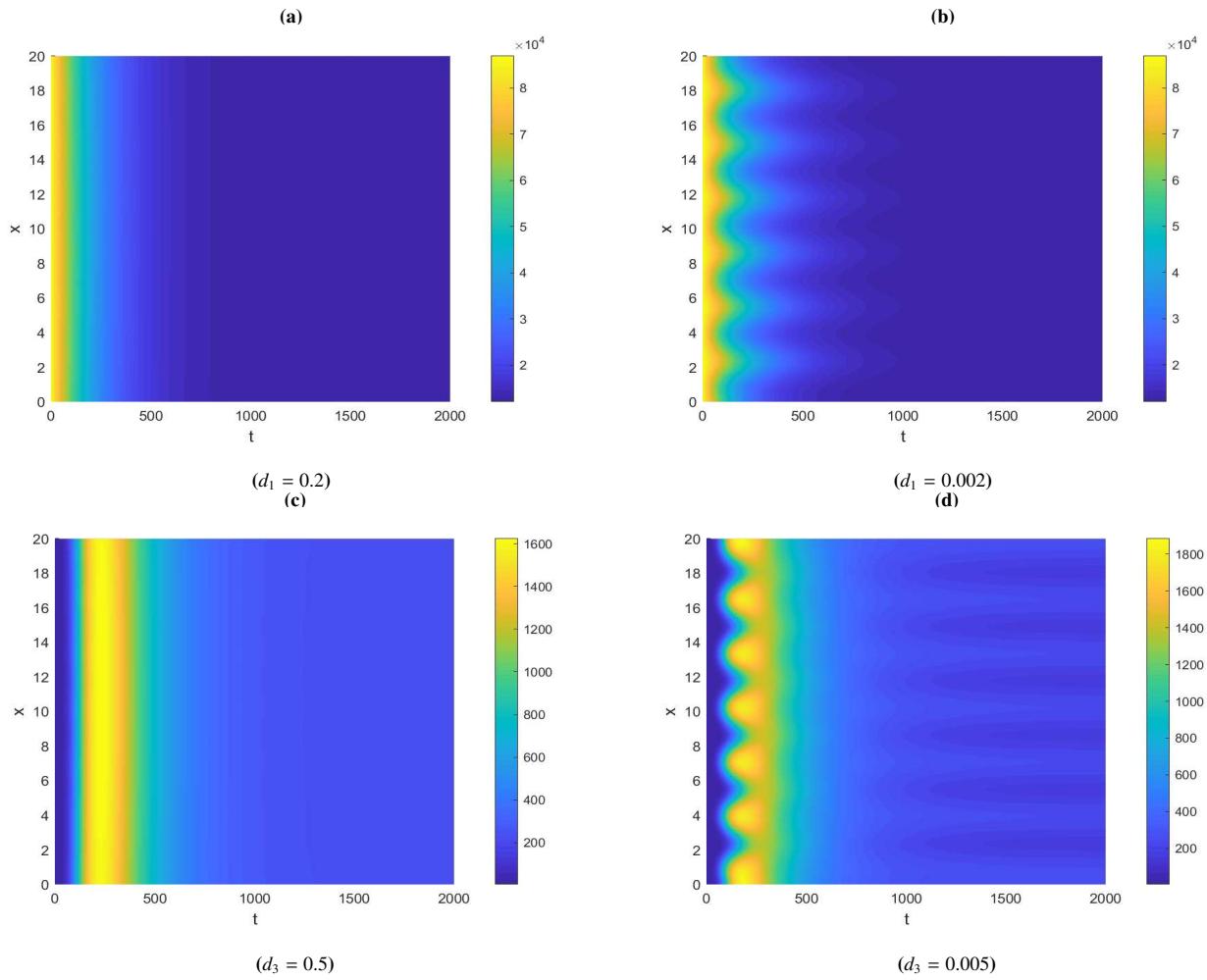


Figure 5. The impact of diffusion coefficients on model (2.1) illness propagation, (a) and (b): d_1 on $S(x, 1500)$, $I(x, 1500)$; (c) and (d): d_3 on $S(x, 1500)$, $I(x, 1500)$.

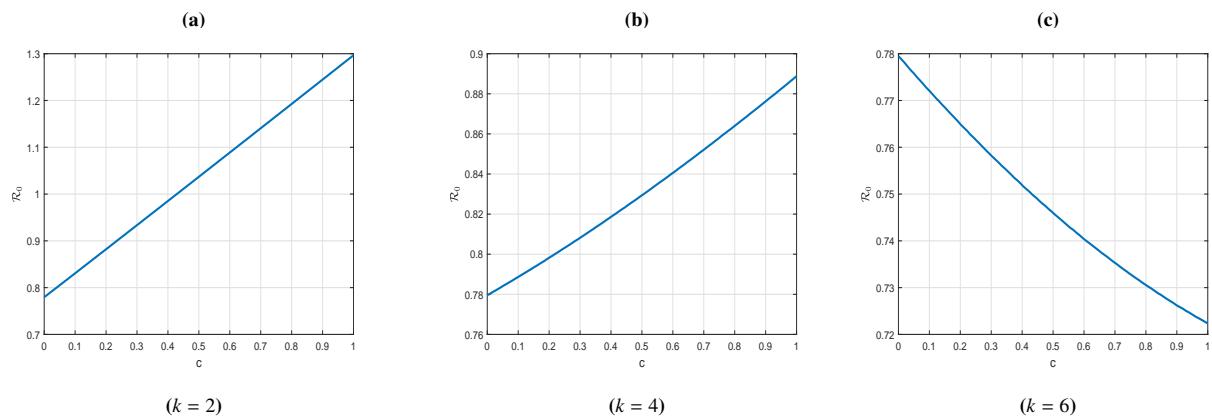


Figure 6. \mathcal{R}_0 in connection with the spatial heterogeneity parameters, $\beta_1 = 1.5 \times 10^{-9} + 1.5 \times 10^{-9}c \sin kx$ and $\beta_2 = 1.88 \times 10^{-9} + 1.88 \times 10^{-9}c \sin kx$, (a) $k = 2$; (b) $k = 4$; (c) $k = 6$.

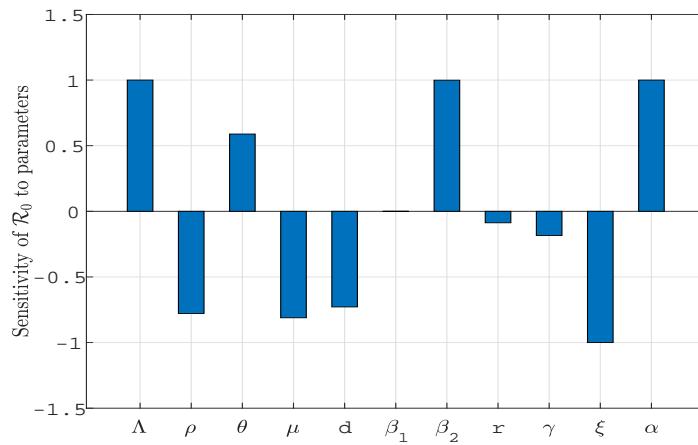


Figure 7. Sensitivity of \mathcal{R}_0 to major parameters.

7. Conclusions

Within this paper, we presented and discussed a SVIR-W spatially heterogeneous model of Cholera that combines multiple transmission pathways, incomplete immunity, general incidence, and Holling II treatment. It turns out the basic reproduction number \mathcal{R}_0 , which is a criterion condition, decided the persistence or extinction of epidemics. In other words, the disease-free steady state \mathcal{E}_0 is globally asymptotically stable with zero maximum treatment rate in case $\mathcal{R}_0 < 1$ (see Theorem 4.2); the illness will be persistent in case $\mathcal{R}_0 > 1$ (see Theorem 4.2). Furthermore, we performed a branching analysis with constant mortality due to disease as a branching parameter (see Theorem 5.1). It can be inferred that the forward branching is always undergone at $\mathcal{R}_0 = 1$, and the presence of positive steady state is entirely excluded as \mathcal{R}_0 is smaller than 1. This means that \mathcal{R}_0 completely determines the persistence and extinction of diseases, which also implies that we can eliminate the disease by controlling parameters such as recruitment rate, bacterial shedding rate, and the spread of environmental viruses to susceptible individuals.

Numerically, we simulated the influence of some crucial parameters on the spatio-temporal distribution of the disease, which is helpful to prevent and manage the disease. Specifically, spatially heterogeneous parameters can cause the disease distribution to show geographical variability (see Figure 4(a)–(l)). The evolution of the dispersal coefficients will affect the spatial distribution of the illness, e.g., as the diffusion coefficient d_3 increases, an infected individual's distribution will quickly become homogeneous (see Figure 5(c) and (d)). We also explored the relationship between the propagation rates $\beta_1(x)$, $\beta_2(x)$, and \mathcal{R}_0 . We note that the monotonicity of the basic reproduction numbers \mathcal{R}_0 and c changes for different values of k (see Figure 6(b) and (c)), which also suggests that spatial heterogeneity dilutes or amplifies the spread of the illness.

It's unfortunate that we only proved the global asymptotic stability of disease-free steady state \mathcal{E}_0 at a maximum treatment rate of $\gamma(x) = 0$. While we have not derived the kinetic behavior of the disease at a maximum treatment rate of $\gamma(x) \neq 0$, we will study this issue in-depth in the future. As is well known, many environmentally spread diseases have incubation periods during which the host can move randomly [46, 47]. That implies that the effect of infection not only relies on the correlation of the present time and location,

but also on the correlation of the previous position, which can generally be characterized by a nonlocal morbidity with a core function. Therefore, it appears relevant and essential to introduce nonlocal effects into models with environmental propagation. This is the focus of our future research.

Use of AI tools declaration

The authors declare they have not used Artificial Intelligence (AI) tools in the creation of this article.

Acknowledgements

The authors would like to thank the anonymous reviewers and the editors for their valuable suggestions for the improvement of the paper. This research is partially supported by the National Natural Science Foundation of Xinjing Uygur Autonomous Region (Grant Nos. 2021D01E12 and 2022TSYCCX0015), the National Natural Science Foundation of China (Grant No. 12361103), the Scientific Research and Innovation Project of Outstanding Doctoral Students in Xinjiang University (Grant No. XJU2022BS022).

Conflict of interest

The authors declare that they have no conflict of interest.

References

1. R. Colwell, A. Huq, Environmental reservoir of *Vibrio cholerae*, the causative agent of cholera, *Ann. Ny. Acad. Sci.*, **740** (1994), 44–53. <https://doi.org/10.1111/j.1749-6632.1994.tb19852.x>
2. A. A. Weil, A. I. Khan, F. Chowdhury, R. C. LaRocque, A. S. G. Faruque, E. T. Ryan, et al., Clinical outcomes of household contacts of patients with cholera in Bangladesh, *Clin. Infect. Dis.*, **49** (2009), 1473–1479. <https://doi.org/10.1086/644779>
3. *World Health Organization, Cholera*. Available from: <https://www.who.int/news-room/fact-sheets/detail/cholera>.
4. *World Health Organization EMRO Yemen cholera situation reports*. Available from: <https://www.emro.who.int/yem/yemeninfocus/situation-reports.html>.
5. V. Rouzier, K. Severe, M. A. Juste, M. Peck, C. Perodin, P. Severe, et al., Cholera vaccination in urban Haiti, *Am. J. Trop. Med. Hyg.*, **89** (2013), 671–681. <https://doi.org/10.4269/ajtmh.13-0171>
6. J. Andrews, S. Basu, Transmission dynamics and control of cholera in Haiti: an epidemic model, *Lancet*, **377** (2011), 1248–1255. [https://doi.org/10.1016/S0140-6736\(11\)60273-0](https://doi.org/10.1016/S0140-6736(11)60273-0)
7. M. C. Eisenberg, Z. S. Shuai, J. H. Tien, P. van den Driessche, A cholera model in a patchy environment with water and human movement, *Math. Biosci.*, **246** (2013), 105–112. <https://doi.org/10.1016/j.mbs.2013.08.003>

8. C. W. Song, R. Xu, A note on the global stability of a multi-strain cholera model with an imperfect vaccine, *Appl. Math. Lett.*, **134** (2022), 108326. <https://doi.org/10.1016/j.aml.2022.108326>
9. X. Y. Zhou, X. Y. Shi, J. A. Cui, Stability and backward bifurcation on a cholera epidemic model with saturated recovery rate, *Math. Method Appl. Sci.*, **40** (2017), 1288–306. <https://doi.org/10.1002/mma.4053>
10. D. H. He, X. Y. Wang, D. Z. Gao, J. Wang, Modeling the 2016-2017 Yemen cholera outbreak with the impact of limited medical resources, *J. Theor. Biol.*, **451** (2018), 80–85. <https://doi.org/10.1016/j.jtbi.2018.04.041>
11. C. Y. Yang, J. Wang, On the intrinsic dynamics of bacteria in waterborne infections, *Math. Biosci.*, **296** (2018), 71–81. <https://doi.org/10.1016/j.mbs.2017.12.005>
12. H. M. N. Teytsa, B. Tsanou, S. Bowong, J. Lubuma, Coupling the modeling of phage-bacteria interaction and cholera epidemiological model with and without optimal control, *J. Theor. Biol.*, **512** (2021), 110537. <https://doi.org/10.1016/j.jtbi.2020.110537>
13. J. Z. Lin, X. Rui, X. H. Tian, Transmission dynamics of cholera with hyperinfectious and hypoinfectious vibrios: mathematical modelling and control strategies, *Math. Biosci. Eng.*, **16** (2019), 4339–4358. <http://dx.doi.org/10.3934/mbe.2019216>
14. J. Y. Yang, C. Modnak, J. Wang, Dynamical analysis and optimal control simulation for an age-structured cholera transmission model, *J. Franklin I.*, **356** (2019), 8438–8467. <https://doi.org/10.1016/j.jfranklin.2019.08.016>
15. J. H. Tien, D. J. D. Earn, Multiple transmission pathways and disease dynamics in a waterborne pathogen model, *B. Math. Biol.*, **72** (2010), 1506–1533. <https://doi.org/10.1007/s11538-010-9507-6>
16. Y. Shi, J.G. Gao, J.L. Wang, Analysis of a reaction-diffusion host-pathogen model with horizontal transmission, *J. Math. Anal. Appl.*, **481** (2020), 123481. <https://doi.org/10.1016/j.jmaa.2019.123481>
17. F. Capone, V. De Cataldis, R. De Luca, Influence of diffusion on the stability of equilibria in a reaction-diffusion system modeling cholera dynamic, *J. Math. Biol.*, **71** (2015), 1107–1131. <https://doi.org/10.1007/s00285-014-0849-9>
18. E. Avila-Vales, Á. G. C. Pérez, Dynamics of a reaction-diffusion SIRS model with general incidence rate in a heterogeneous environment, *Z. Angew. Math. Phys.*, **73** (2022), 1–23. <https://doi.org/10.1007/s00033-021-01645-0>
19. F. B. Wang, J. P. Shi, X. F. Zou, Dynamics of a host-pathogen system on a bounded spatial domain, *Commun. Pur. Appl. Anal.*, **14** (2015), 2535–2560. <https://doi.org/10.3934/cpaa.2015.14.2535>
20. E. Avila-Vales, G. E. Garcia-Almeida, Á. G. C. Pérez, Qualitative analysis of a diffusive SIR epidemic model with saturated incidence rate in a heterogeneous environment, *J. Math. Anal. Appl.*, **503** (2021), 125295. <https://doi.org/10.1016/j.jmaa.2021.125295>
21. J. L. Wang, F. L. Xie, T. Kuniya, Analysis of a reaction-diffusion cholera epidemic model in a spatially heterogeneous environment, *Commun. Nonlinear Sci.*, **80**, (2020), 104951. <https://doi.org/10.1016/j.cnsns.2019.104951>

22. X. D. Chen, R. H. Cui, Global stability in a diffusive cholera epidemic model with nonlinear incidence, *Appl. Math. Lett.*, **111** (2021), 106596. <https://doi.org/10.1016/j.aml.2020.106596>

23. Y. Yang, L. Zou, J. L. Zhou, C. H. Hsu, Dynamics of a waterborne pathogen model with spatial heterogeneity and general incidence rate, *Nonlinear Anal.-Real.*, **53** (2020), 103065. <https://doi.org/10.1016/j.nonrwa.2019.103065>

24. X. Y. Wang, F. B. Wang, Impact of bacterial hyperinfectivity on cholera epidemics in a spatially heterogeneous environment, *J. Math. Anal. Appl.*, **480** (2019), 123407. <https://doi.org/10.1016/j.jmaa.2019.123407>

25. J. L. Wang, X. Q. Wu, Dynamics and profiles of a diffusive cholera model with bacterial hyperinfectivity and distinct dispersal rates, *J. Dyn. Differ. Equations*, **35** (2023), 1205–1241. <http://dx.doi.org/10.1007/s10884-021-09975-3>

26. D. M. Hartley, J. G. Morris Jr, D. L. Smith, Hyperinfectivity: a critiquelement in the ability of *V.cholerae* to cause epidemics?, *Plos Med.*, **3** (2006), e7. <https://doi.org/10.1371/journal.pmed.0030007>

27. H. L. Smith, *Monotone Dynamical Systems: An Introduction to the Theory of Competitive and Cooperative Systems*, American Mathematical Society, Providence, 1995. <https://doi.org/10.1090/surv/041>

28. H. L. Smith, X. Q. Zhao, Robust persistence for semidynamical systems, *Nonlinear Anal.*, **47** (2001), 6169–6179. [https://doi.org/10.1016/S0362-546X\(01\)00678-2](https://doi.org/10.1016/S0362-546X(01)00678-2)

29. Y. J. Lou, X. Q. Zhao, A reaction-diffusion malaria model with incubation period in the vector population, *J. Math. Biol.*, **62** (2011), 543–568. <https://doi.org/10.1007/s00285-010-0346-8>

30. Y. X. Wu, X. F. Zou, Dynamics and profiles of a diffusive host-pathogen system with distinct dispersal rates, *J. Differ. Equations*, **264** (2018), 4989–5024. <https://doi.org/10.1016/j.jde.2017.12.027>

31. H. Y. Cheng, Y. F. Lv, R. Yuan, Long time behavior of a degenerate NPZ model with spatial heterogeneity, *Appl. Math. Lett.*, **132** (2022), 108088. <https://doi.org/10.1016/j.aml.2022.108088>

32. S. B. Hsu, F. B. Wang, X. Q. Zhao, Dynamics of a periodically pulsed bio-reactor model with a hydraulic storage zone, *J. Dyn. Differ. Equations*, **23** (2011), 817–842. <https://doi.org/10.1007/s10884-011-9224-3>

33. G. Sell, Y. You, *Dynamics of Evolutionary Equations*, Springer, New York, 2002. <https://doi.org/10.1007/978-1-4757-5037-9>

34. P. Magal, X. Q. Zhao, Global attractors and steady states for uniformly persistent dynamical systems, *SIAM J. Math. Anal.*, **37** (2005), 251–275. <https://doi.org/10.1137/S0036141003439173>

35. X. Q. Zhao, *Dynamics Systems in Population Biology*, Springer-Verlag, New York, 2017. <https://doi.org/10.1007/978-0-387-21761-1>

36. W. D. Wang, X. Q. Zhao, Basic reproduction numbers for reaction-diffusion epidemic models, *SIAM J. Appl. Dyn. Syst.*, **11** (2012), 1652–1673. <https://doi.org/10.1137/120872942>

37. H. R. Thieme, Spectral bound and reproduction number for infinite-dimensional population structure and time heterogeneity, *SIAM J. Appl. Math.*, **70** (2009), 188–211. <https://doi.org/10.1137/080732870>

38. K. J. Engel, R. Nagel, *One-Parameter Semigroups for Linear Evolution Equations*, Springer, New York, 1999. <https://doi.org/10.1007/b97696>

39. R. D. Nussbaum, Eigenvectors of nonlinear positive operator and the linear Krein-Rutman theorem, in *Fixed Point Theory*, Springer, (1981), 309–331. <https://doi.org/10.1007/bfb0092191>

40. R. H. Martin, H. L. Smith, Abstract functional differential equations and reaction-diffusion systems, *T. Am. Math. Soc.*, **321** (1990), 1–44. <https://doi.org/10.1090/s0002-9947-1990-0967316-x>

41. Y. Jin, F. B. Wang, Dynamics of a benthic-drift model for two competitive species, *J. Math. Anal. Appl.*, **462** (2018), 840–860. <https://doi.org/10.1016/j.jmaa.2017.12.050>

42. W. M. Ni, *The Mathematics of Diffusion*, Society for Industrial and Applied Mathematics, Philadelphia, 2011. <https://doi.org/10.1137/1.9781611971972>

43. J. P. Shi, X. F. Wang, On global bifurcation for quasilinear elliptic systems on bounded domains, *J. Differ. Equations*, **246** (2009), 2788–2812. <https://doi.org/10.1016/j.jde.2008.09.009>

44. M. G. Crandall, P. H. Rabinowitz, Bifurcation from simple eigenvalues, *J. Funct. Anal.*, **8** (1971), 321–340. [https://doi.org/10.1016/0022-1236\(71\)90015-2](https://doi.org/10.1016/0022-1236(71)90015-2)

45. J. P. Shi, Persistence and bifurcation of degenerate solutions, *J. Funct. Anal.*, **69** (1999), 494–531. <https://doi.org/10.1006/jfan.1999.3483>

46. H. Y. Shu, Z. M. Ma, X. S. Wang, Threshold dynamics of a nonlocal and delayed cholera model in a spatially heterogeneous environment, *J. Math. Biol.*, **83** (2021), 1–33. <https://doi.org/10.1007/s00285-021-01672-5>

47. J. X. Xu, J. L. Wang, Threshold-type result for a nonlocal diffusive cholera model with seasonally forced intrinsic incubation period, *Discrete Cont. Dyn.-B*, **28** (2023), 3393–3413. <https://doi.org/10.3934/dcdsb.2022223>

48. Y. H. Grad, J. C. Miller, M. Lipsitch, Cholera modeling: challenges to quantitative analysis and predicting the impact of interventions, *Epidemiology*, **23** (2012), 523–530. <https://doi.org/10.1097/EDE.0b013e3182572581>

49. X. N. Wang, H. Wang, M. Y. Li, \mathcal{R}_0 and sensitivity analysis of a predator-prey model with seasonality and maturation delay, *Math. Biosci.*, **315** (2019), 108225. <https://doi.org/10.1016/j.mbs.2019.108225>



AIMS Press

© 2024 the Author(s), licensee AIMS Press. This is an open access article distributed under the terms of the Creative Commons Attribution License (<http://creativecommons.org/licenses/by/4.0>)