

# Localized outbreaks in S-I-R model with diffusion

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We investigate an SIRS epidemic model with spatial diffusion and nonlinear incidence rates. We show that for small diffusion rate of the infected class  $D_I$ , the infected population tends to be highly localized at certain points inside the domain, forming  $K$  spikes. We then study three distinct destabilization mechanisms, as well as a transition from localized spikes to plateau solutions. Two of the instabilities are due to coarsening (spike death) and self-replication (spike birth), and have well-known analogues in other reaction-diffusion systems such as the Schnakenberg model. The third transition is when a single spike becomes unstable and moves to the boundary. This happens when the diffusion of the recovered class,  $D_R$  becomes sufficiently small. In all cases, the stability thresholds are computed asymptotically and are verified by numerical experiments. We also show that the spike solution can transit into an plateau-type solution when the diffusion rates of recovered and susceptible class are sufficiently small. Implications for disease spread and control through quarantine are discussed.

## 1. INTRODUCTION

The SIRS epidemic model introduced by Kermack and McKendrick in 1927 [1] is widely used to model the spread of infectious diseases. The population is divided into three disjoint classes: susceptible (S), infected (I), and recovered (R), where susceptibles can be infected by those already infected and subsequently recover, and recovered class are immune to the disease but lose immunity over time. These assumptions are modelled using the following system of ODEs:

$$\begin{cases} S_t = -\beta SI + \gamma R, \\ I_t = \beta SI - \nu I, \\ R_t = \nu I - \gamma R, \end{cases} \quad (1)$$

where  $\beta$  is the infection rate,  $\nu$  is the recovery rate, and  $\gamma$  is the rate of immunity loss. Kermack and McKendrick's work has motivated the use of mathematics in the study of epidemiology [2–4].

While spatially-homogeneous dynamics are by now well studied, modelling spatial interactions is still an active area of research. Most disease outbreaks have a strong spatial characteristic, and many studies emphasize the importance of the spatial dimension for modelling these outbreaks. For example, [5] looked at spatio-temporal patterns in HIV outbreaks in Malawi over two decades (1994–2010). The authors found that the disease initially spread in several localized hot-spots and they identified several geographically differentiated HIV/AIDS epidemics rather than a single one. These initial outbreaks were followed by a complex spatio-temporal dynamics. Similar spatial clusters of HIV outbreaks were found in a recent study [6] in Phayao Province, Thailand, and in South Africa [7].

In recent decades, numerous methodologies have been used to describe spatial distribution of disease. This includes the use of cellular automata [8, 9], metapopulations [10–12], networks [13, 14] and partial differential equations [15, 16]. Generally speaking, incorporating spatial structure leads to very rich dynamics in epidemic models, such as formation of disease hot-spots.

In this paper we study spatially-localized outbreaks for the SIRS model with spatial dispersion. As will be shown below, such outbreaks can occur when the infection rate  $\beta$  is nonlinear. For simplicity, we will assume that  $\beta$  is proportional to  $I$ , although other types of nonlinearity, such as Holling functional response also lead to hot-spot formation. We model spatial dispersion using diffusion. This results in the following system,

$$\begin{cases} S_t = D_S S_{xx} - \chi SI^2 + \gamma R, \\ I_t = D_I I_{xx} + \chi SI^2 - \nu I, \\ R_t = D_R R_{xx} + \nu I - \gamma R. \end{cases} \quad (2)$$

Here  $D_S, D_I, D_R$  are diffusion coefficients of each class of population,  $\chi I$  is the rate of infection. We study the epidemic system on 1-D interval  $[-L, L]$  with Neumann boundary conditions, so that  $S_x = I_x = R_x = 0$  at  $x = \pm L$ . For simplicity, we also assume the timescale of infection and recovery is much shorter than the average life span, so birth and death rates for each class are neglected.

The second key assumption we make is that the infected class  $I$  diffuses more slowly than others. There are two scenarios where this is biologically plausible. The first scenario, common in many species is that the disease itself reduces the species mobility. A second scenario, applicable to humans, is an intentional quarantine policy to limit the spread of infection. Such a policy is well known to be effective in controlling disease outbreaks and is often used as a first-line defense against quickly-spreading infections.

We therefore write  $D_I = \varepsilon^2$  where  $\varepsilon$  is small. By further rescaling [17], we may set  $\chi = 1, \nu = 1$ . This leads to the following singularly perturbed reaction diffusion system:

$$\begin{cases} S_t = D_S S_{xx} - SI^2 + \gamma R, \\ I_t = \varepsilon^2 I_{xx} + SI^2 - I, \\ R_t = D_R R_{xx} + I - \gamma R. \end{cases} \quad (3)$$

Under these assumptions, this system has localized disease concentrations corresponding to spike-type solutions. Such spike patterns have been studied in great detail since 1990's in simpler reaction-diffusion systems consisting of two components, such as Gierer-Meinhardt system, Gray-Scott model, Schnakenberg model and Keller-Segel model and its variants. We refer reader to [18–30] and references therein. The introduction of a third component leads to interesting new phenomena not present in two-component reaction-diffusion systems [31].

Let us summarize the main results in this paper. Simulations and analysis show that the behavior of the system is highly dependent on diffusion rate  $D_R$ , relative to the diffusion rate of infected class,  $\varepsilon^2$ . We isolate two distinct regimes: either  $D_R \gg O(\varepsilon^2)$  or  $D_R \leq O(\varepsilon^2)$ .

The regime  $D_R \gg O(\varepsilon^2)$  is studied in Sections 2, 3, 5. In this regime, the steady-state population consists of  $K$  hot-spots of disease, uniformly distributed inside the interval  $[-L, L]$ . Depending on system parameters, the  $K$ -spike steady state can undergo two types of instabilities. The first type, analyzed in Section 5.1 is referred to as *spike competition instability*. As a result of such an instability, some of the hot-spots are “absorbed” by others, resulting in fewer hot-spots. The second type of instability, studied in 5.2 is referred to as *self-replication instability*, whereby a spike splits into two, resulting in more spikes. These instabilities are illustrated in Figure 1. Figure 1(a) shows 8 spikes that gradually coarsen into 2 as  $D_S$  is gradually increased. On the other hand, with one-spike equilibrium as initial condition, replication occurs and more spikes appear as we gradually decrease  $D_S$ . This is shown in Fig 1(b). We derive explicit thresholds for  $D_S$  such that the spike competition occurs when  $D_S > D_{SK}^{com}$ ,  $K \geq 2$ ; and self-replication instability occurs when  $D_S < D_{SK}^{rep}$ ,  $K \geq 1$ . Formulas for  $D_{SK}^{com}$  and  $D_{SK}^{rep}$  are given in Section 5.

The second regime we study is when  $D_R$  is small:  $D_R \leq O(\varepsilon^2)$ . In this case, a single spike can become unstable, and depending on other parameters, two phenomena can occur. If  $D_R$  is sufficiently small, a single spike moves to the boundary (depending on how big  $D_S$  is), as illustrated in Figure 2. This phenomenon is studied in Section 4. On the other hand, when both  $D_R$  and  $D_S$  are small, the spike “fattens up” and becomes a mesa-type pattern, i.e. a contiguous region of high concentration of disease connected via a sharp interface to a region of low concentration. Numerically we observe two types of inhomogeneous equilibrium depending on the value of  $D_S$  and an example of such a steady state pattern is shown in Figure 3. Spike-type solution exists for sufficiently large  $D_S$ , but transition to interface-type patterns for small  $D_S$ . This process is illustrated in Figure 3(left). Interface patterns are studied in Section 6.

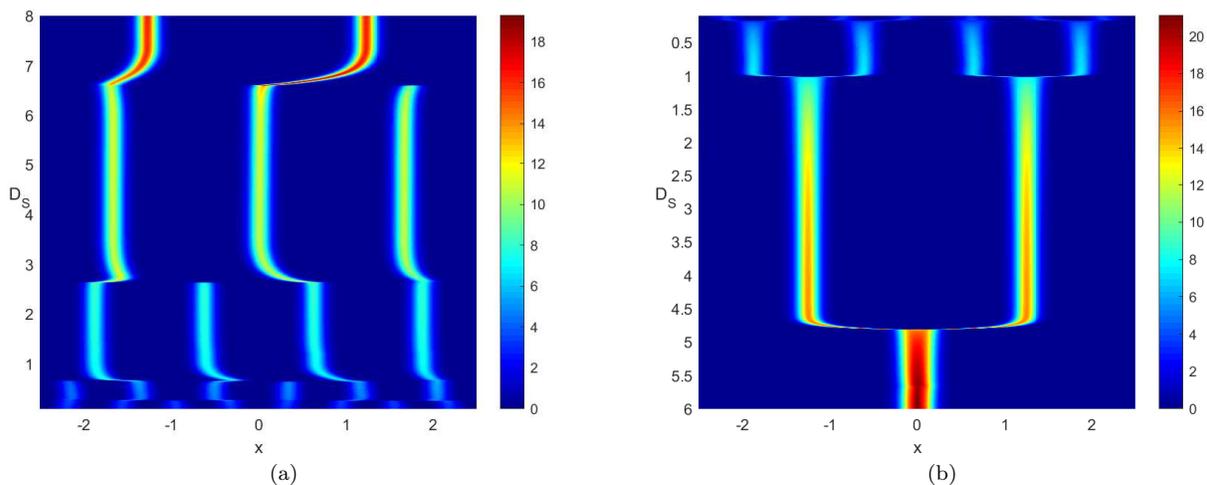


FIG. 1. Instabilities of steady state spike solutions induced by slowly increasing  $D_S$  or decreasing  $D_S$ . Here  $D_R = 1, L = 2.5, \varepsilon = 0.05, N = 15$  and  $\gamma = 1$ . Left: Coarsening (competition) instability when  $D_S$  is increased ( $D_s = 1 + 10^{-5}t$ ). Colour plot of  $I$  is shown. Right: Self-replication instability when  $D_S$  is slowly decreased ( $D_s = 6 - 10^{-5}t$ ).

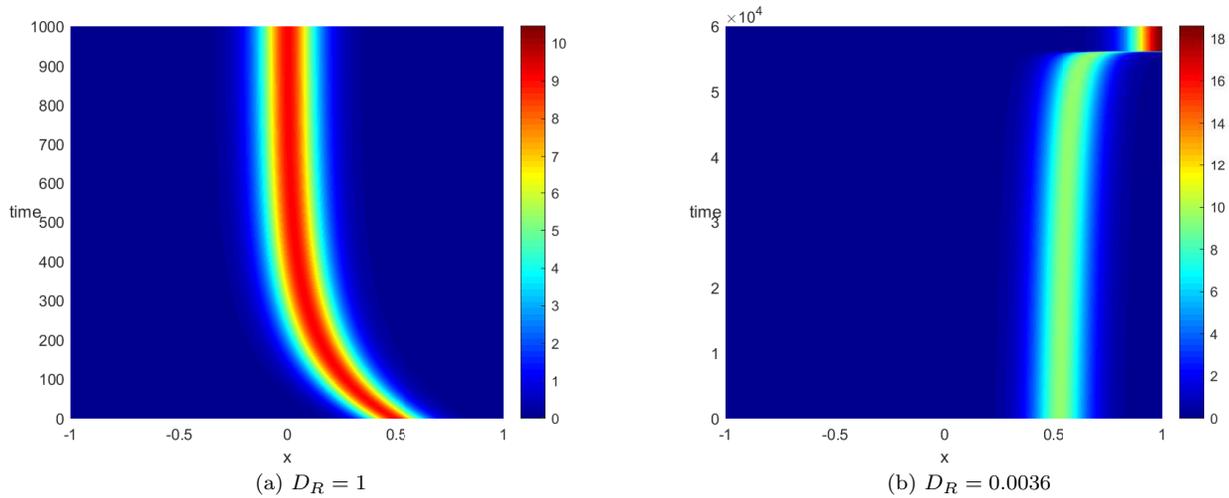


FIG. 2. Stable and unstable motion of a single spike. Here,  $D_R$  is as indicated while other parameters are fixed at  $D_S = 5$ ,  $L = 1$ ,  $\varepsilon = 0.06$ ,  $N = 5$  and  $\gamma = 1$ . In (a), one-spike equilibrium moves to the center, which shows that the center spike is stable. In (b) the spike moves to boundary instead of moving to center, so that a single spike is unstable.

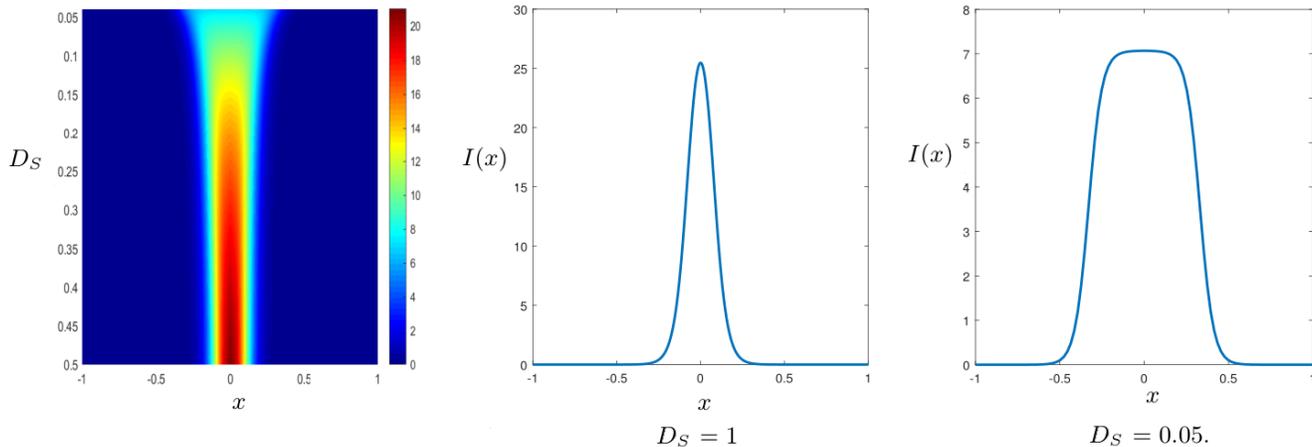


FIG. 3. Transition from spike to mesa when  $D_R = 0$ ,  $\varepsilon = 0.04$ ,  $N = 10$ ,  $L = 1$ ,  $\gamma = 1$  with  $D_S$  as a control parameter. Left:  $D_S$  is gradually decreased from 0.5 to 0.05. Middle, Right: Profile of  $I(x)$  for  $D_S$  as indicated.

## 2. SINGLE-SPIKE SOLUTION

We start by constructing a single interior spike solution to (3). Such a solution corresponds to a localized concentration of the infected population  $I$  at some point  $x_0$  in the interior of the domain,  $x_0 \in (-L, L)$ . The extent of the spike is of  $O(\varepsilon)$ . We therefore introduce the inner variable

$$y = \frac{x - x_0}{\varepsilon}. \quad (4)$$

In the inner region, equilibrium solution of (3) then becomes

$$\begin{cases} S_{yy} - \frac{\varepsilon^2}{D_S} S I^2 + \frac{\varepsilon^2}{D_S} \gamma R = 0, \\ I_{yy} + S I^2 - I = 0, \\ R_{yy} + \frac{\varepsilon^2}{D_R} I - \frac{\varepsilon^2}{D_R} \gamma R = 0. \end{cases} \quad (5)$$

We then expand  $S, I, R$  as

$$\begin{aligned} S &= S_0 + \varepsilon S_1 + O(\varepsilon^2), \\ I &= I_0 + \varepsilon I_1 + O(\varepsilon^2), \\ R &= R_0 + \varepsilon R_1 + O(\varepsilon^2). \end{aligned} \quad (6)$$

Upon substituting (6) into (5) and collecting higher-order terms in  $\varepsilon$ , we obtain, to leading order,

$$\begin{cases} S_{0yy} = 0, \\ I_{0yy} + S_0 I_0^2 - I_0 = 0, \\ R_{0yy} = 0. \end{cases} \quad (7)$$

This shows that  $S_0$  and  $R_0$  are constants to be determined. We then rescale

$$I_0 = \frac{1}{S_0} w(y), \quad (8)$$

where so that  $w$  satisfies the well-known ground-state

$$w'' - w + w^2 = 0, \quad w \rightarrow 0 \text{ as } y \rightarrow \pm\infty$$

whose explicit solution is given by

$$w(y) = \frac{3}{2} \operatorname{sech}^2\left(\frac{y}{2}\right).$$

To determine  $R_0$  and  $S_0$ , we must match the inner and the outer region. In the outer region we approximate  $I \sim \left(\int_{-L}^L I dx\right) \delta(x - x_0)$  and  $SI^2 \sim \left(\int_{-L}^L SI^2 dx\right) \delta(x - x_0)$ . We further estimate  $\left(\int_{-L}^L I dx\right) \sim \frac{1}{S_0} \varepsilon \int_{-\infty}^{\infty} w dy \sim 6\varepsilon/S_0$  and similarly,  $\int_{-L}^L SI^2 dx \sim 6\varepsilon/S_0$ , so that

$$\begin{aligned} 0 &= D_S S_{xx} + \gamma R - 6\varepsilon/S_0 \delta(x - x_0), \\ 0 &= D_R R_{xx} - \gamma R + 6\varepsilon/S_0 \delta(x - x_0). \end{aligned} \quad (9)$$

To solve (9), we introduce the modified Green's function  $G(x; x_0)$ , which satisfies

$$\begin{cases} G_{xx} - \frac{\gamma}{D_R} G = -\delta(x; x_0), \\ G_x(x_0^+) - G_x(x_0^-) = -1, \\ G_x(\pm L) = 0. \end{cases} \quad (10)$$

A simple calculation gives

$$G(x; x_0) = \begin{cases} \frac{\cosh\left(\sqrt{\frac{\gamma}{D_R}}(x+L)\right) \cosh\left(\sqrt{\frac{\gamma}{D_R}}(x_0-L)\right)}{\sqrt{\frac{\gamma}{D_R}} \sinh\left(2\sqrt{\frac{\gamma}{D_R}}L\right)}, & -L < x < x_0 \\ \frac{\cosh\left(\sqrt{\frac{\gamma}{D_R}}(x_0+L)\right) \cosh\left(\sqrt{\frac{\gamma}{D_R}}(x-L)\right)}{\sqrt{\frac{\gamma}{D_R}} \sinh\left(2\sqrt{\frac{\gamma}{D_R}}L\right)}, & x_0 < x < L. \end{cases} \quad (11)$$

The solution to (9) is then given by

$$R(x) = \frac{6\varepsilon}{S_0 D_R} G(x; x_0) \quad (12)$$

and

$$S(x) = -\frac{D_R}{D_S} R(x) + \frac{D_R}{D_S} R_0 + S_0, \quad (13)$$

where  $R_0 = \frac{6\varepsilon}{S_0 D_R} G(x; x_0)$  and  $S_0$  is to be determined.

To find  $S_0$ , we use the conservation of mass. Let  $N$  be the total population, so that

$$N \equiv \int_{-L}^L S + I + R dx. \quad (14)$$

Note that by adding three equations in (3) and integrating over the domain,  $N$  is independent of time. We will also take

$$N = 2N_0L, \quad (15)$$

where  $N_0$  is an arbitrary constant depending on initial conditions, so that  $N$  scales with domain size;  $N_0$  can be thought of an average density.

We now substitute (8), (12) and (13) into the mass conservation condition (14, 15) to obtain that

$$S_0^2 - N_0S_0 + E = 0, \quad (16)$$

where

$$E = \frac{3\varepsilon}{L} \left( 1 + \frac{1}{\gamma} - \frac{D_R}{\gamma D_S} + 2\sqrt{\frac{D_R}{\gamma}} \frac{L}{D_S} \frac{\cosh(\sqrt{\frac{\gamma}{D_R}}(x_0 + L)) \cosh(\sqrt{\frac{\gamma}{D_R}}(x_0 - L))}{\sinh(2\sqrt{\frac{\gamma}{D_R}}L)} \right). \quad (17a)$$

Solving (16) we get two roots when  $N_0^2 - 4E > 0$ . Asymptotically in  $\varepsilon$ , they are

$$S_{0-} \sim \frac{E}{N_0}, \quad (17b)$$

which is of  $O(\varepsilon)$  and

$$S_{0+} \sim N_0, \quad (17c)$$

which is of  $O(1)$ . Plots of these two roots are shown in Fig 4. The two roots connect at a fold point corresponding to a double root of (16).

We now summarize our first result:

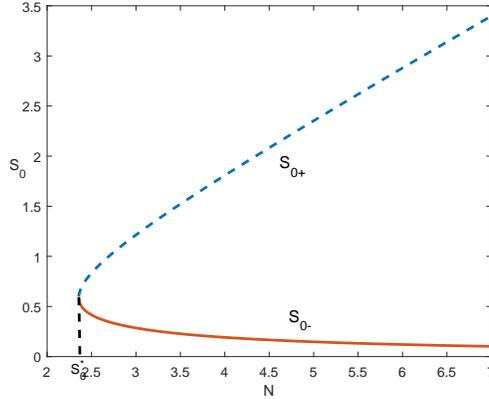


FIG. 4. Plot of two roots to (16) vs. total population  $N$ . Here  $\varepsilon = 0.05$ ,  $L = 1$ ,  $D_R = 1$ ,  $D_S = 1$  and  $\gamma = 1$ . The dashed curve denotes  $S_{0+}$  and the solid curve denotes  $S_{0-}$ .  $S_0^*$  is the fold point where (16) has double root.

**Result 2.1** *With  $D_I = \varepsilon^2$  and  $0 < \varepsilon \ll 1$ , the SIRS system (3) has the following single-spike steady state:*

$$\begin{aligned} S(x) &\sim -\frac{6\varepsilon}{S_0 D_S} G(x; x_0) + \frac{6\varepsilon}{S_0 D_S} G(x_0; x_0) + S_0, \\ I(x) &\sim \frac{1}{S_0} w\left(\frac{x - x_0}{\varepsilon}\right), \\ R(x) &\sim \frac{6\varepsilon}{S_0 D_R} G(x; x_0), \end{aligned} \quad (18)$$

where  $G(x; x_0)$  is given by (11),  $w(y) = \frac{3}{2} \operatorname{sech}^2(\frac{y}{2})$  and  $S_0$  is a constant determined by the total population mass as given in (17).

### 3. NONLOCAL EIGENVALUE PROBLEM

We now study the stability of one-spike solution. We first linearize around the steady state by taking

$$\begin{aligned} S(x, t) &= S(x) + e^{\lambda t} \varphi(x), \\ I(x, t) &= I(x) + e^{\lambda t} \psi(x), \\ R(x, t) &= R(x) + e^{\lambda t} \xi(x). \end{aligned}$$

Assuming  $|\varphi|, |\psi|, |\xi| \ll O(1)$  we obtain the linearized problem

$$\begin{cases} \lambda \varphi = D_S \varphi_{xx} - I_0^2 \varphi - 2S_0 I_0 \psi + \gamma \xi, \\ \lambda \psi = \varepsilon^2 \psi_{xx} + I_0^2 \varphi + (2S_0 I_0 - 1) \psi, \\ \lambda \xi = D_R \xi_{xx} + \psi - \gamma \xi. \end{cases} \quad (19)$$

In the inner region, we let  $y = \frac{x-x_0}{\varepsilon}$  where  $x_0$  is the spike position. To leading order, we then obtain  $\varphi_{yy} \sim 0$  so that  $\varphi(x) \sim \varphi_0$  is constant to be determined. The equation for  $\psi$  is

$$\lambda \psi = \psi_{yy} - \psi + 2w(y)\psi + I_0^2 \varphi_0. \quad (20)$$

In the outer region, we approximate

$$\begin{cases} D_S \varphi_{xx} - \lambda \varphi + \gamma \xi = c_1 \delta(x; x_0), \\ D_R \xi_{xx} - (\gamma + \lambda) \xi = c_2 \delta(x; x_0). \end{cases} \quad (21)$$

where

$$c_1 = \left( \varphi_0 \int I_0^2 dx + 2S_0 \int I_0 \psi dx \right), \quad c_2 = - \int \psi dx. \quad (22)$$

We write

$$\xi(x; x_0) = -\frac{c_2}{D_R} G \left( x; x_0, \sqrt{\frac{\gamma + \lambda}{D_R}} \right), \quad (23)$$

where  $G(x; x_0, \mu)$  is the Green's function that satisfies

$$\begin{cases} G_{xx} - \mu^2 G = -\delta(x; x_0), \\ G_x(\pm L) = 0, \end{cases} \quad (24)$$

and is explicitly given by

$$G = \frac{1}{\mu \sinh(2\mu L)} \begin{cases} \cosh(\mu(x+L)) \cosh(\mu(x_0-L)), & -L < x < x_0 \\ \cosh(\mu(x_0+L)) \cosh(\mu(x-L)), & x_0 < x < L. \end{cases} \quad (25)$$

To solve for  $\varphi$ , we make a change of variables. Let

$$\varphi = \frac{D_R \gamma}{\lambda(D_R - D_S) - \gamma D_S} \xi + \varphi_h. \quad (26)$$

Then  $\varphi_h$  satisfies:

$$D_S \varphi_{hxx} - \lambda \varphi_h = \left( c_1 - \frac{\gamma c_2}{\lambda(D_R - D_S) - \gamma D_S} \right) \delta(x; x_0)$$

so that

$$\varphi_h = -\frac{1}{D_S} \left( c_1 - \frac{\gamma c_2}{\lambda(D_R - D_S) - \gamma D_S} \right) G \left( x; x_0; \sqrt{\frac{\lambda}{D_S}} \right)$$

Therefore we estimate

$$\varphi_0 = \varphi(x_0) \sim -c_2 \frac{\gamma}{\lambda(D_R - D_S) - \gamma D_S} G \left( x_0; x_0; \sqrt{\frac{\gamma + \lambda}{D_R}} \right) - \frac{1}{D_S} \left( c_1 - \frac{\gamma c_2}{\lambda(D_R - D_S) - \gamma D_S} \right) G \left( x_0; x_0; \sqrt{\frac{\lambda}{D_S}} \right)$$

and

$$c_1 = \varepsilon \left( \frac{\varphi_0}{S_0^2} \int w^2 dy + 2 \int w\psi dy \right); \quad c_2 = -\varepsilon \int \psi dy. \quad (27)$$

After some algebra, this leads to the following non-local eigenvalue problem (NLEP),

$$(L_0 - \lambda)\psi = w^2 \frac{2}{\int_{-\infty}^{\infty} w^2 dy - (\lambda + 1) \frac{S_0^2}{\varepsilon P}} \int_{-\infty}^{\infty} w\psi dy, \quad (28)$$

$$\text{where } L_0\psi = \psi_{yy} - \psi + 2w\psi \quad (29)$$

and where

$$P = \frac{\gamma \sqrt{\frac{D_R}{\lambda + \gamma}} \cosh\left(\sqrt{\frac{\lambda + \gamma}{D_R}}(x_0 + L)\right) \cosh\left(\sqrt{\frac{\lambda + \gamma}{D_R}}(x_0 - L)\right)}{\lambda(D_R - D_S) - \gamma D_S \sinh\left(2\sqrt{\frac{\lambda + \gamma}{D_R}}L\right)} - \frac{\lambda + 1 + \frac{D_S \gamma}{\lambda(D_R - D_S) - \gamma D_S} \cosh\left(\sqrt{\frac{\gamma}{D_S}}(x_0 + L)\right) \cosh\left(\sqrt{\frac{\gamma}{D_S}}(x_0 - L)\right)}{\sqrt{\lambda D_S} \sinh\left(2\sqrt{\frac{\gamma}{D_S}}L\right)}. \quad (30)$$

For the special case when  $x_0 = 0$ , this expression simplifies to

$$P(\lambda) = \frac{\gamma \sqrt{\frac{D_R}{\lambda + \gamma}} \coth\left(\sqrt{\frac{\lambda + \gamma}{D_R}}L\right)}{\lambda(D_R - D_S) - \gamma D_S} - \frac{\lambda + 1 + \frac{D_S \gamma}{\lambda(D_R - D_S) - \gamma D_S} \coth\left(\sqrt{\frac{\gamma}{D_S}}L\right)}{\sqrt{\lambda D_S}}. \quad (31)$$

In general, the NLEP problem (28) is difficult to tackle since  $P$  has such a complicated dependence on  $\lambda$ . However there are two cases for which stability of (28) is well established: namely, large or small  $\frac{S_0^2}{\varepsilon}$ . Note that  $S_0$  is given by (17) and has two branches,  $S_{0+}$  and  $S_{0-}$ , refer to Figure 4. Consider the case of large  $N$ . Then  $\frac{S_{0+}^2}{\varepsilon} \gg 1$  whereas  $\frac{S_{0-}^2}{\varepsilon} \ll 1$ . In the former case, (28) reduces to a *local* eigenvalue problem  $(L_0 - \lambda)\psi \sim 0$ . This problem is well known to admit a positive eigenvalue  $\lambda = 5/4$  so that this branch is unstable. For the latter case ( $S_0 = S_{0-}$ ), the problem (28) reduces to the following well-known NLEP problem:

$$\lambda\psi = L_0\psi - 2w^2 \frac{\int_{-\infty}^{\infty} w\psi dy}{\int_{-\infty}^{\infty} w^2 dy}, \quad S_0 \sim O(\varepsilon). \quad (32)$$

This is well-known to be *stable* as was first proven in [19].

Finally, a lengthy but a straightforward algebraic computation shows that at the fold point where  $S_{0+} = S_{0-}$ , there is a zero eigenvalue whose corresponding eigenfunction is given by  $\psi = w$ . This suggests that the entire branch  $S_{0+}$  is unstable whereas the entire branch  $S_{0-}$  is stable, although the proof of this fact is not in the cards due to the complex structure of  $P(\lambda)$ . This structure is analogous to the well-known properties of the Grey-Scott model in the low-feed regime [32].

Another approach is to consider the limit of large  $D_R$  and/or  $D_S$  (various so-called shadow limits). We do not observe any additional instabilities of a single spike in this regime so we will not pursue it further here.

#### 4. SPIKE MOTION.

We now study the motion of the interior spike, which is determined by small eigenvalues. We rewrite the system as following:

$$\begin{cases} S_t = D_S S_{xx} - SI^2 + \gamma R, \\ I_t = \varepsilon^2 I_{xx} + SI^2 - I, \\ R_t = D_R R_{xx} + I - \gamma R \end{cases} \quad (33)$$

with Neumann boundary conditions and  $D_S, D_R \gg O(\varepsilon^2)$ . To study the motion of the spike, we expand around the center  $x_0$  by writing  $x = x_0 + \varepsilon y$ , and let  $x_0 = x_0(\varepsilon^2 t)$ ,  $S(y, t) = S\left(\frac{x - x_0(\varepsilon^2 t)}{\varepsilon}\right)$ ,  $I(y, t) = I\left(\frac{x - x_0(\varepsilon^2 t)}{\varepsilon}\right)$ , and

$R(y, t) = R\left(\frac{x-x_0(\varepsilon^2 t)}{\varepsilon}\right)$ . Then system (33) becomes

$$\begin{cases} -\varepsilon^3 x'_0 S_y = D_S S_{yy} - \varepsilon^2 S I^2 + \varepsilon^2 \gamma R, \\ -\varepsilon x'_0 I_y = I_{yy} + S I^2 - I, \\ -\varepsilon^3 x'_0 R_y = D_R R_{yy} + \varepsilon^2 I - \varepsilon^2 \gamma R. \end{cases} \quad (34)$$

Applying the same expansion (6) and collecting  $\varepsilon$  order, we obtain that

$$\begin{cases} S_{1yy} = 0, \\ -x'_0 I_{0y} = I_{1yy} + \chi S_1 I_0^2 + 2\chi S_0 I_0 I_1 - I_1, \\ R_{1yy} = 0, \end{cases} \quad (35)$$

in which  $S_0, I_0, R_0$  are expressed in (17b), (12) and (13). Multiply the second equation by  $I_{0y}$  and integrate to obtain the solvability condition

$$x'_0 \int_{-\infty}^{\infty} I_{0y}^2 dy = \frac{1}{3} \int_{-\infty}^{\infty} I_0^3 S_{1y} dy. \quad (36)$$

From (35), we know that  $S_1$  is linear so that  $S_{1y}$  is a constant. To determine  $S_{1y}$ , we match to the outer region. We expand

$$\begin{aligned} S(x; x_0) &= S(x_0 + \varepsilon y; x_0) \\ &= S(x_0) + \varepsilon y S'(x_0), \end{aligned} \quad (37)$$

where  $S(x; x_0)$  in outer region is expressed in (12). We then match it with the expansion (6) to have

$$S_1 = \left( -\frac{6\varepsilon}{S_0 D_S} G'(x_0; x_0) \right) y, \quad (38)$$

where  $G(x; x_0)$  is expressed in (11). Therefore we have

$$S_{1y} = -\frac{6\varepsilon}{S_0 D_S} \begin{cases} G(x_0^+; x_0), & -L < x < x_0 \\ G(x_0^-; x_0), & x_0 < x < L. \end{cases} \quad (39)$$

Substituting (39) into the equation (36) gives the equation that describes the motion of the interior spike:

$$x'_0 = -\frac{6\varepsilon}{D_S S_0^2} \frac{\sinh\left(2\sqrt{\frac{\gamma}{D_R}} x_0\right)}{\sinh\left(2\sqrt{\frac{\gamma}{D_R}} L\right)}, \quad (40)$$

where  $S_0$  is expressed in (17b). Write

$$S_0 = \varepsilon \hat{S}, \quad (41)$$

then we have

$$\frac{dx_0}{dt} = -\frac{6\varepsilon}{D_S \hat{S}^2} \frac{\sinh\left(2\sqrt{\frac{\gamma}{D_R}} x_0\right)}{\sinh\left(2\sqrt{\frac{\gamma}{D_R}} L\right)}. \quad (42)$$

It's obvious to see that equation (42) has one equilibrium  $x_0 = 0$ , and the corresponding eigenvalue is

$$\lambda = -\frac{12\varepsilon}{D_S \hat{S}^2} \frac{\sqrt{\frac{\gamma}{D_R}}}{\sinh\left(2\sqrt{\frac{\gamma}{D_R}} L\right)} < 0. \quad (43)$$

Therefore the equilibrium centered at  $x_0 = 0$  is stable with respect to spike motion.

#### 4.1. Boundary effects and spike motion

From formula (43), it is clear that the eigenvalue is *stable*, provided that  $D_R$  is not too small. However numerical experiments show that the spike becomes unstable and moves to the boundary when  $D_R$  is of  $O(\varepsilon^2)$ . To understand this, note that for small  $D_R$ , (17b) simplifies to

$$S_0 \sim \frac{6\varepsilon(1 + \frac{1}{\gamma})}{N} \quad (44)$$

and  $\hat{S} \sim \frac{6(1+\frac{1}{\gamma})}{N}$ . Therefore (43) simplifies to

$$\lambda \sim -\frac{2}{3} \frac{\varepsilon N^2}{D_S(1 + \frac{1}{\gamma})^2} \sqrt{\frac{\gamma}{D_R}} \exp\left(-2L\sqrt{\frac{\gamma}{D_R}}\right), \quad D_R \ll 1. \quad (45)$$

As such, the effect of  $\langle S_x \rangle$  becomes exponentially small. On the other hand, there are also exponentially weak boundary effects due to the interaction of the pulse with the boundary that we disregarded in the computation leading to (36). These boundary terms appear when integrating by parts in (36). To compute them, we replace (36) by a more precise expression

$$-x'_0 \int I_{0y}^2 dy = (I_{0y}I_{1y} - I_0I_1)|_{y=\frac{-L-x_0}{\varepsilon}}^{y=\frac{L-x_0}{\varepsilon}} + \int I_{0y}I_0^2 S_1 dy. \quad (46)$$

The computation of the boundary terms is relatively standard and we summarize it here. Note that

$$w(y) \sim 6e^{-y} \text{ as } y \rightarrow \infty \quad (47)$$

so that

$$I_0 \sim \frac{6}{S_0} \exp(-y). \quad (48)$$

For  $x$  near  $L$ , we change variables:

$$x = L + \varepsilon z$$

so that  $y = \frac{L-x_0}{\varepsilon} + z$  and

$$I_0 \sim \frac{6}{S_0} \exp\left(-\frac{L-x_0}{\varepsilon}\right) \exp(z). \quad (49)$$

Near  $z = 0$ , equation for  $I_1$  satisfies  $I_{1zz} - I_1 \sim 0$ , so that  $I_1 = A \exp z + B \exp(-z)$ . Since  $I'(L) = 0$ , we must therefore have

$$I_1 \sim \frac{6\varepsilon}{S_0} \exp\left(-\frac{L-x_0}{\varepsilon}\right) \exp(-z), \quad (50)$$

so that

$$(I_{0x}I_{1x} - I_0I_1)|_{x=L} = -\frac{72\varepsilon}{S_0^2} \exp\left(-2\frac{L-x_0}{\varepsilon}\right). \quad (51)$$

Performing a similar computation at  $x = -L$ , and evaluating the remaining terms as before, we obtain

$$x'_0 \sim -\frac{6\varepsilon}{D_S} \frac{\sinh\left(2\sqrt{\frac{\gamma}{D_R}}x_0\right)}{\sinh\left(2\sqrt{\frac{\gamma}{D_R}}L\right)} \frac{1}{\hat{S}^2} + 60\varepsilon \left\{ \exp\left(2\frac{x_0-L}{\varepsilon}\right) - \exp\left(2\frac{-L-x_0}{\varepsilon}\right) \right\} \quad (52)$$

so that

$$\lambda \sim -\frac{12\varepsilon}{D_S} \frac{\sqrt{\frac{\gamma}{D_R}}}{\sinh\left(2\sqrt{\frac{\gamma}{D_R}}L\right)} \frac{1}{\hat{S}^2} + 240 \exp\left(-\frac{2L}{\varepsilon}\right). \quad (53)$$

This expression clearly shows that the boundary term can play a destabilizing effect when the first term on the right hand side of (53) is exponentially small. This happens precisely when  $D_R$  is small. Setting  $\lambda = 0$ , substituting  $\hat{S} \sim \frac{6(1+\frac{1}{\gamma})}{N}$  and solving for  $D_s$  yields the critical value

$$D_S^* \sim \frac{\varepsilon N^2}{1440} \sqrt{\frac{\gamma}{D_R}} \exp\left(2L\left(\frac{1}{\varepsilon} - \sqrt{\frac{\gamma}{D_R}}\right)\right) \quad (54)$$

with a single spike centered at center being unstable when  $D_S > D_S^*$ , and stable otherwise. This phenomenon is illustrated in Figure 3. Take  $L = 2, \gamma = 1, N = 5$  and  $D_R = 0.005298$ . Then (54) yields  $D_S^* = 5.00$ . It follows that a single spike is unstable at the origin when  $D_R < 0.005298$  and is stable otherwise. This is confirmed in Figure 3.

The boundary effect discussed here is similar to destabilization discussed in [33]. However the difference here is that this effect is primarily driven by having a small  $D_R$ , and is very specific to having three components. Although the asymptotics of (54) are valid as long as  $D_R \gg O(\varepsilon^2)$ , similar destabilization phenomenon also happens when  $D_R = O(\varepsilon^2)$ . Although the asymptotics of (54) break down in such a case, numerics show that the destabilization phenomenon persists as  $D_S$  is increased.

## 5. INSTABILITY THRESHOLDS OF MULTI-SPIKE EQUILIBRIUM

In this section we study  $K$ -spike patterns, where  $K \geq 1$ . We analyze two types of instabilities, one is referred to as spike competition or coarsening instability, whereby some of the spikes are annihilated if the initial state contains too many spikes. The other is referred to as self replication, whereby a new spike may appear by the process of spike splitting. In this chapter we derive explicit thresholds for these instabilities.

### 5.1. Coarsening

When there are too many spikes, some of them get absorbed by others. This is known as coarsening or competition instability. To determine the instability threshold for spike competition, we apply the method in [34, 35] and compute the critical value at which an asymmetric spike pattern bifurcates from symmetric branch. To do this, consider a single interior spike on the domain  $[-l, l]$ . Duplicating the domain  $K$  times we obtain  $K$  spikes on the domain of size  $2L = 2lK$ . From (13) we have:

$$S(l) = S_0 + \frac{3\varepsilon}{D_S S_0} \sqrt{\frac{D_R}{\gamma}} \left( \frac{1}{\tanh\left(\sqrt{\frac{\gamma}{D_R}} l\right)} - \frac{1}{\sinh\left(\sqrt{\frac{\gamma}{D_R}} l\right)} \right), \quad (55)$$

where  $S_0 = S_{0-}$  is given in (17b) with  $x_0 = 0$ . Plots of  $S(l)$  when  $D_S = 1$  and  $D_S = 3$  are shown in Fig 5. The bifurcation point corresponds to the minimum point of the curve  $l \rightarrow S(l)$ . Setting  $S'(l) = 0$  then yields the critical stability threshold. Solving for  $D_S$  as a function of other parameters, and upon substituting  $l = L/K$  we obtain the critical threshold

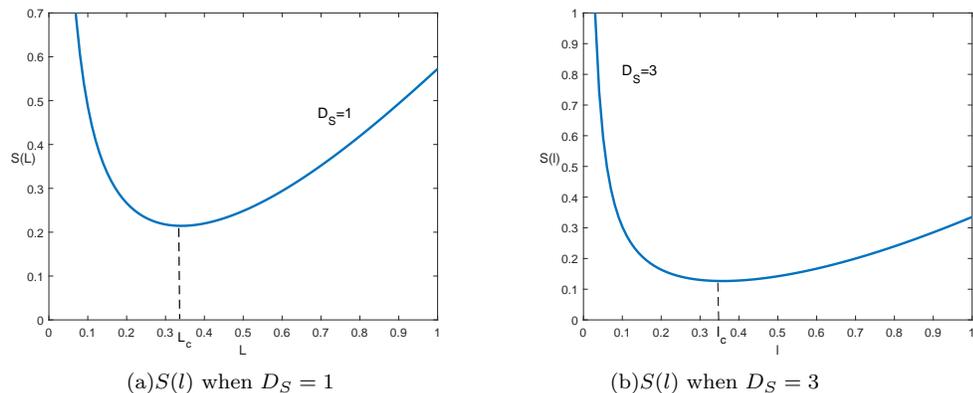


FIG. 5. Plots of function  $S(l)$  versus  $l$  for  $D_S = 1$  and  $D_S = 3$ . Other parameters are fixed and they are:  $\varepsilon = 0.02, D_R = 2, \gamma = 1, N_0 = 4$ .

$$\begin{aligned}
D_{SK}^{com} \sim & \frac{N_0^2 \left(\frac{L}{K}\right)^3}{3\varepsilon \left(1 + \frac{1}{\gamma}\right)^2} \left( 1 - \frac{1}{\tanh^2 \left(\sqrt{\frac{\gamma}{D_R} \frac{L}{K}}\right)} + \frac{\cosh \left(\sqrt{\frac{\gamma}{D_R} \frac{L}{K}}\right)}{\sinh^2 \left(\sqrt{\frac{\gamma}{D_R} \frac{L}{K}}\right)} \right) \\
& + \frac{N_0^2 \left(\frac{L}{K}\right)^2}{3\varepsilon \left(1 + \frac{1}{\gamma}\right)^2} \sqrt{\frac{D_R}{\gamma}} \left( \frac{1}{\tanh \left(\sqrt{\frac{\gamma}{D_R} \frac{L}{K}}\right)} - \frac{1}{\sinh \left(\sqrt{\frac{\gamma}{D_R} \frac{L}{K}}\right)} \right).
\end{aligned} \tag{56}$$

The  $K$ -spike solution is unstable and some of the spikes will disappear when  $D_S > D_{SK}^{com}$ . The plot of  $D_{SK}^{com}$  as a function of  $D_R$  is shown in Figure 6. Note that  $D_{SK}^{com}$  has the following asymptotics as  $D_R \rightarrow \infty$ :

$$D_{SK}^{com} \sim \frac{N_0^2 \left(\frac{L}{K}\right)^3}{3\varepsilon \left(1 + \frac{1}{\gamma}\right)^2}, \quad \text{as } D_R \rightarrow \infty, \tag{57}$$

which is shown in Fig 6. We now summarize the following result:

**Result 5.1** Consider a  $K$ -spike solution for the system (3) on an interval of length  $2L$  with  $K > 1$ . Then in the limit of  $\varepsilon \rightarrow 0$ , this solution is stable provided that  $D_S < D_{SK}^{com}$ , where  $D_{SK}^{com}$  is given by (56). When  $D_S > D_{SK}^{com}$ , the  $K$ -spike solution becomes unstable due to competition (or coarsening) instability and some of the spikes disappear.

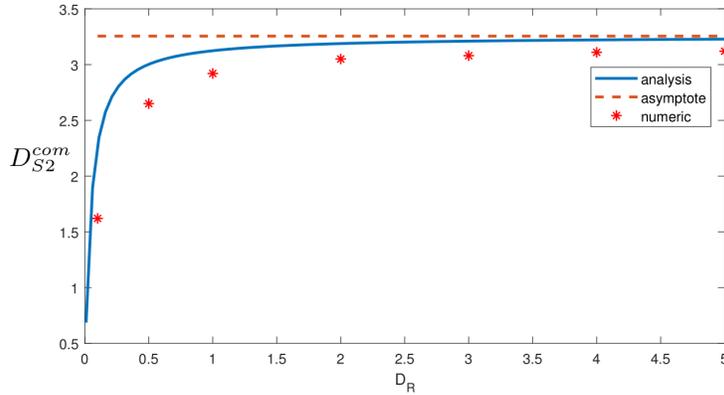


FIG. 6. Plot of stability threshold  $D_{S2}^{com}$  vs.  $D_R$  for single spike solutions. Here  $\varepsilon = 0.02$ ,  $L = 1$ ,  $N_0 = 2.5$ , and  $\gamma = 1$ . The curve denotes analysis value  $D_{S2}^{com}$  obtained by (56), and the dashed line is the asymptote of the curve. The dots are obtained by numeric simulations, and they have a good agreement with analysis.

## 5.2. Self-replication

Unlike coarsening instability, self-replication is related to disappearance of the single spike equilibrium solution. The mechanism has been studied in detail for Gray-Scott model [22, 23, 36–39], and it is similar here. We start by changing variables

$$S(x) = \frac{\varepsilon}{\sqrt{D_S}} \tilde{S}(x), \quad I(x) = \frac{\sqrt{D_S}}{\varepsilon} \tilde{I}(x), \quad x = \varepsilon y, \tag{58}$$

so that the system (3) transforms to

$$\begin{cases} \frac{\sqrt{D_S}}{\varepsilon} \tilde{S}_{yy} - \frac{\sqrt{D_S}}{\varepsilon} \tilde{S} \tilde{I}^2 + \gamma R = 0, \\ \tilde{I}_{yy} + \tilde{S} \tilde{I}^2 - \tilde{I} = 0, \\ \frac{D_R}{\varepsilon^2} R_{yy} + \frac{\sqrt{D_S}}{\varepsilon} \tilde{I} - \gamma R = 0. \end{cases} \tag{59}$$

Next, assume that  $D_S, D_R$  are  $O(1)$ . Then to leading order, in the inner region we obtain the following problem, referred to as the **core problem**,

$$\begin{cases} \tilde{S}_{yy} - \tilde{S}\tilde{I}^2 = 0, \\ \tilde{I}_{yy} + \tilde{S}\tilde{I}^2 - \tilde{I} = 0, \end{cases} \quad (60a)$$

this core problem is identical to the core problem for both the Grey-Scott model [22, 23, 36, 39], and the Schnakenberg model [27]. Assuming that the spike is symmetric, we define

$$A := \tilde{S}_y(\infty) = \int_0^\infty \tilde{S}\tilde{I}^2 dy. \quad (60b)$$

By plotting the numerical bifurcation diagram of (60b), it was found in [23, 39] that the steady state disappears when  $A > A_c \approx 1.35$ , and this disappearance leads to self-replication. To determine  $A$  in terms of the other parameters of the problem, we perform an asymptotic matching to the outer region. We estimate  $\int SI^2 = \int I$  and

$$D_R R_{xx} - \gamma R = - \left( \int I dx \right) \delta(x), \quad D_S S_{xx} + \gamma R = - \left( \int I dx \right) \delta(x).$$

The solution is then given by

$$R(x) = \frac{\left( \int I dx \right)}{D_R} G(x), \quad (61a)$$

$$S(x) = -\frac{D_R}{D_S} R(x) + \frac{D_R}{D_S} R(0). \quad (61b)$$

where as before,

$$G(x) = \frac{\sqrt{\frac{D_R}{\gamma}}}{2 \sinh\left(\sqrt{\frac{\gamma}{D_R}} l\right)} \begin{cases} \cosh\left(\sqrt{\frac{\gamma}{D_R}}(x+l)\right), & -l < x < 0 \\ \cosh\left(\sqrt{\frac{\gamma}{D_R}}(x-l)\right), & x < x < l. \end{cases} \quad (61c)$$

We substitute (61) into the total mass equation (14) to obtain that

$$N = 2N_0 l = 2A\sqrt{D_S} \left(1 + \frac{1}{\gamma} - \frac{D_R}{\gamma D_S}\right) + \frac{2Al}{\sqrt{D_S}} \frac{\sqrt{\frac{D_R}{\gamma}}}{\tanh\left(\frac{\gamma}{D_R} l\right)}, \quad (62)$$

so that self-replication occurs when

$$A = \frac{N_0 l \sqrt{D_S}}{D_S \left(1 + \frac{1}{\gamma}\right) + l \frac{\sqrt{\frac{D_R}{\gamma}}}{\tanh\left(\sqrt{\frac{\gamma}{D_R}} l\right)} - \frac{D_R}{\gamma}} > A_c \approx 1.35. \quad (63)$$

Equivalently, we rewrite (63) to obtain the following quadratic equation with respect to  $\sqrt{D_S}$

$$\left(1 + \frac{1}{\gamma}\right) D_S - \frac{N_0 l}{A_c} \sqrt{D_S} + \frac{\sqrt{\frac{D_R}{\gamma}} l}{\tanh\left(\sqrt{\frac{\gamma}{D_R}} l\right)} - \frac{D_R}{\gamma} = 0. \quad (64)$$

Therefore replication of one-spike solution occurs when  $D_S < D_S^{rep}$ , where  $\sqrt{D_S^{rep}}$  is the large root of (64). For  $K$  spikes on domain  $[-L, L]$  with  $L = Kl$ , this leads to the following result:

**Result 5.2** Consider a  $K$ -spike solution of the system (3) on an interval of length  $2L$  with  $K \geq 1$ . Then in the limit of  $\varepsilon \rightarrow 0$ , this solution is stable provided that  $D_S > D_{SK}^{rep}$ , where  $D_S = D_{SK}^{rep}$  is the root of (64), in which  $l = \frac{L}{K}$  and  $A_c \approx 1.35$  corresponds to the fold point of the problem (60).

Figure 7 shows numerical validation of Result 5.2. The solid curve denotes the asymptotic curve as given in Result 5.2. Above the curve, a single spike is stable. As  $D_S$  is decreased and crosses the curve, self-replication takes place resulting in two spikes. The dots denote numeric simulations. Good agreement is observed between numerics and asymptotics.

For a fixed  $D_R$  and a given number of spikes  $K$ , we have derived both upper and lower thresholds on the  $D_S$  for which  $K$  spikes are stable. Note that multiple solutions (e.g. two or three spikes) can be stable at the same time. This is illustrated in Figure 8.

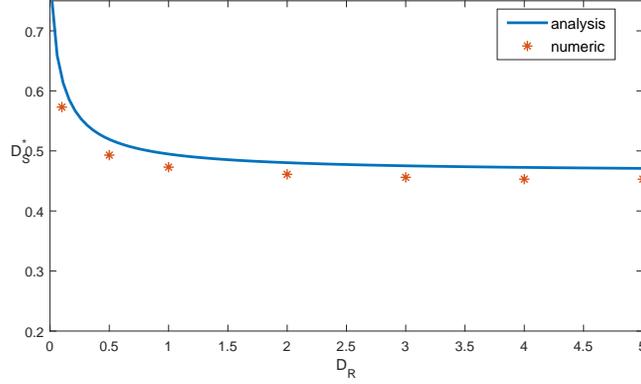


FIG. 7. Self-replication threshold (Result 5.2). Comparison between numerics and analysis. Solid curve is the analytical result given by (64). Dots denote numerical simulations. Self-replication is observed as  $D_S$  is decreased past the solid curve in the figure. Here  $\varepsilon = 0.005$ ,  $N = 5$ ,  $L = 1$ , and  $\gamma = 1$ .

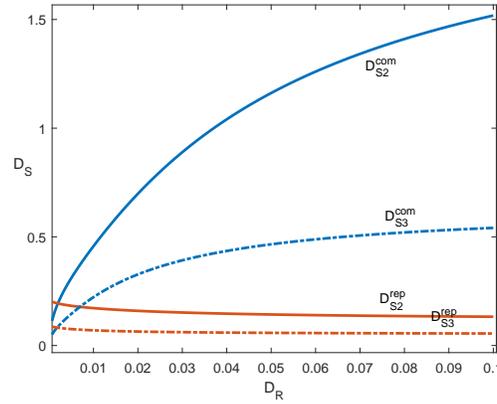


FIG. 8. Bifurcation diagram of  $K$ -spike patterns for  $K = 2, 3$ . The region between solid curves is the stable region for 2-spike patterns, and the region between dashed curves is the stable region for 3-spike patterns. Above the regions spike competition instability occurs, below the region, self replication instability occurs. Here  $\varepsilon = 0.03$ ,  $N_0 = 2.5$ ,  $L = 1$ ,  $\gamma = 1$ .

## 6. MESA-LIKE STEADY STATES WHEN $D_R = 0$ .

As shown in Section 5.5.1, multi-spike configurations lose stability when  $D_R$  is sufficiently small: even a single spike eventually becomes unstable (due to an exponentially small eigenvalue becoming positive) and moves towards the boundary when  $D_R = O(\varepsilon^2)$ . For even smaller values of  $D_R$ , we observe numerically that the spike “fattens” as shown in Figure 9. In the limit of  $D_R \rightarrow 0$ , numerics indicate a phase separation of infected population. This can be thought of as a “quarantene effect”: when mobility of recovered population and susceptible population is reduced, the infected population is confined to a certain region of the entire domain with a sharp interface inbetween.

Here we perform the analysis for the limiting case  $D_R = 0$  and  $D_S$  being small, although similar results hold even when  $D_R = D_S$  and is nonzero. At the steady state, we then have  $I = \gamma R$  so that the model (3) reduces to

$$\begin{cases} 0 = D_S S_{xx} - SI^2 + I, \\ 0 = \varepsilon^2 I_{xx} + SI^2 - I. \end{cases} \quad (65)$$

Adding the two equations we we obtain that  $D_S S + \varepsilon^2 I$  is constant. We then eliminate  $S$  from the second equation to obtain

$$D_S I_{xx} = \frac{D_S}{\varepsilon^2} I - CI^2 + I^3, \quad (66)$$

where

$$C = I + \frac{D_S}{\varepsilon^2} S \quad (67)$$

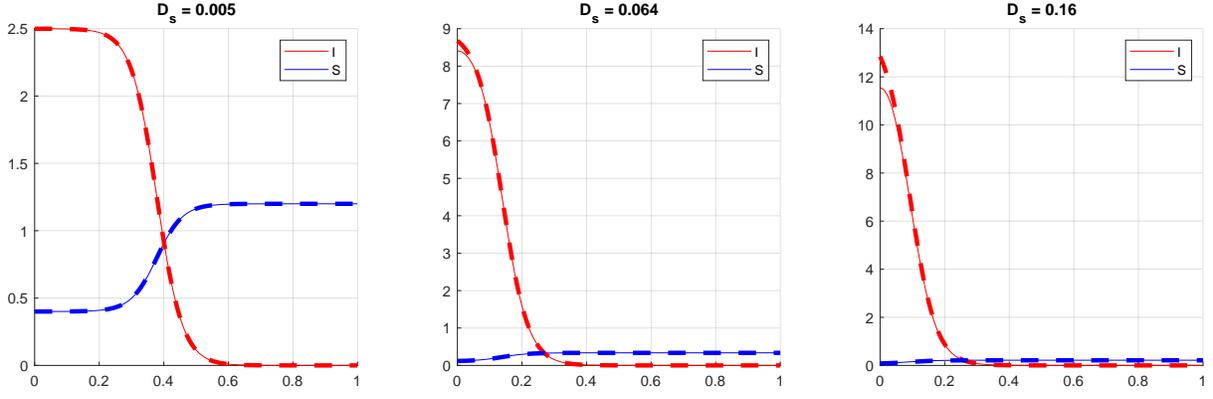


FIG. 9. Steady states of the system (3) with  $L = 1, D_R = 0, \varepsilon = 0.04, \gamma = 1, N_0 = 2.79$  and with  $D_S$  as indicated. Solid curves correspond to the full numerical solution of (3). Dashed lines show the asymptotic approximation (69).

is a constant to be determined.

Equation (66) admits a heteroclinic solution connecting the steady state  $I = 0$  to a positive steady state  $I_+$  provided that the Maxwell-line condition holds:  $\int_{I_0}^{I_+} \left( \frac{D_S}{\varepsilon^2} I - CI^2 + I^3 \right) dI = 0$ . This is equivalent to cubic having equidistant roots, that is,

$$\frac{D_S}{\varepsilon^2} I - CI^2 + I^3 = I \left( I - \frac{I_+}{2} \right) (I - I_+) \quad (68)$$

so that

$$I_+ = \sqrt{\frac{2D_S}{\varepsilon^2}}, \quad C = \frac{3}{2} I_+. \quad (69a)$$

In this case there is an interface solution on the domain  $[0, L]$  given by

$$\gamma R = I \sim I_+ \left( \frac{1}{2} \tanh \left( \frac{I_+}{2\sqrt{2}} \frac{(l - |x|)}{\sqrt{D_S}} \right) + \frac{1}{2} \right); \quad (69b)$$

$$S \sim \frac{\varepsilon^2}{D_S} I_+ \left( 1 - \frac{1}{2} \tanh \left( \frac{I_+}{2\sqrt{2}} \frac{(l - |x|)}{\sqrt{D_S}} \right) \right). \quad (69c)$$

Here,  $l$  is the location of the interface. A back-to-back interface solution such as shown in figure 3 is obtained by extending this solution to  $[-L, L]$  using even reflection. Finally, the interface location  $l$  is determined using the mass conservation condition,  $N_0 L = \int_0^L (S + I + R) dx$ . In the limit  $\varepsilon \rightarrow 0$ , this yields

$$LN_0 = I_+ \left\{ \left( 1 + \frac{1}{\gamma} \right) l + \left( \frac{3}{2} L - \frac{1}{2} l \right) \left( \frac{\varepsilon^2}{D} \right) \right\}$$

so that solving for  $l$  we obtain

$$l = \frac{N_0 - I_+ \frac{3}{2} \left( \frac{\varepsilon^2}{D} \right)}{I_+ \left( 1 + \frac{1}{\gamma} - \left( \frac{\varepsilon^2}{D} \right) \right)} L. \quad (69d)$$

This result is valid as long as  $O(\varepsilon) \ll l < L$ . In this case, the interface has an exponentially weak effect on the boundary, and the agreement with the numerics is nearly perfect. Figure illustrates this. Solution (69) is shown super-imposed on the numerical solution; the difference is imperceptible in the “eye-ball norm” as long as  $l = O(1)$ . The asymptotics break down when  $l$  becomes small (figure 9, right), and the interface transforms into a spike solution.

Note that the infected class subdivides the domain into outbreak portion ( $x < l$ ) and disease-free portion ( $x > l$ ). The susceptible population is three times smaller within the outbreak portion of the domain when compared with the disease-free portion.

For simplicity, we took  $D_R = 0$  here. Numerical simulations indicate that similar interface solutions persist for sufficiently small  $D_R$ , although it changes  $l$  as well as the interface shape. We defer their study to future work.

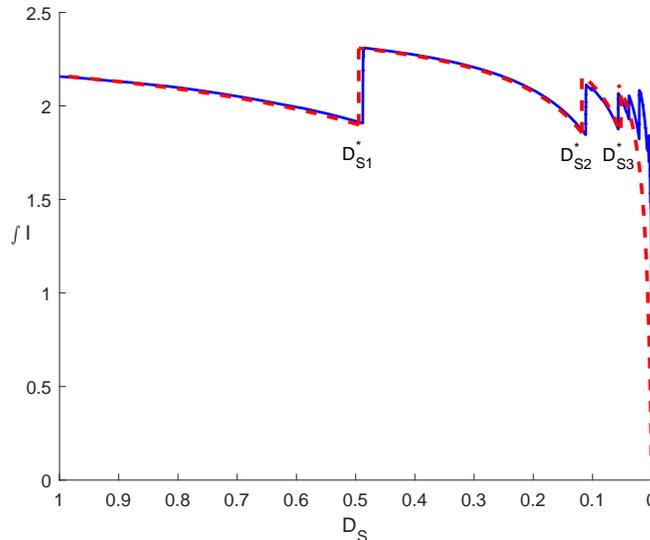


FIG. 10. Total mass of infected people versus  $D_S$ . Here  $\varepsilon = 0.005$ ,  $N = 5$ ,  $D_R = 1$ ,  $L = 1$ , and  $\gamma = 1$ . The solid curve denotes numeric results, and dashed curves is the asymptotic result corresponding to self-replication thresholds of Result 5.2.

## 7. DISCUSSION

In this paper we studied the consequence of adding spatial diffusion to the relatively-standard SIRS model. Under certain reasonable assumptions, the resulting system (3) has a very rich solution space, exhibiting hot-spots as well as interface-type solutions, depending on whether  $D_R$  is large or small, respectively.

Occurrence of disease clusters has been widely documented in epidemiology literature, see e.g. [5–7, 40, 41]. Our study underscores the importance of diffusion in formation of hot-spots and disease spread. One of the key assumptions leading to hot-spot formation was that the diffusion of infected class is relatively slow compared to the susceptible class. While it is difficult to measure (or even quantify) diffusion rates, one study [7] did find a strong positive correlation between HIV hot-spot location and proximity to a large road. A multitude of other causes have been proposed (see [40] and references therein). This includes the level of male circumcision; religiosity (less HIV prevalence in muslim communities in Africa); urbanization level with wider HIV prevalence in rural areas, among others; preponderance of drug use [41].

The hot-spot regime  $D_R \gg D_I$  is very similar to the previous analysis for the two-component reaction-diffusion systems, such as the Schnakenberg model [27, 35], and the behaviour is qualitatively similar to the SI model with diffusion introduced in [16] (which itself is a generalization of the Schnakenberg model). However, from the analysis point of view, the third component introduces a novel non-local eigenvalue problem (see Section 3). On the other hand, the regime  $D_R \leq D_I$  requires completely new analysis. On one hand, the resolution of an exponentially small boundary layer in Section 4.4.1 is crucial for computing stability thresholds of a single interior spike in this regime. On the other, this regime also leads to mesa-type solutions of Section 6. The analysis there is similar to interface solutions derived in [42, 43] for the Gray-Scott model. However it appears to be more robust: such interface solutions exist for a wide range of parameters here, rather than a very narrow range studied in [42, 43].

In Figure 10 we plot the total mass of infected population versus  $D_S$ . As  $D_S$  is decreased, the mobility of susceptible population is reduced and initially leads to a decrease of overall disease load. However as  $D_S$  is decreased further, eventually a self-replication threshold is triggered. This results in an immediate increase of infection hot-spots and an overall increase in infected population. This underscores a highly nonlinear relationship between mobility and disease outbreaks.

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