# ELECTRONIC SUPPLEMENTARY MATERIAL

# SPATIAL EPIDEMICS DYNAMICS: SYNCHRONIZATION

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# CONTENTS

1	Introduction	1
2	Bifurcation Diagram	1
3	Deterministic Solution	2
4	Stochastic Simulation4.1 Gillespie Algorthim	
5	Coherence Dependence on Parameters 5.1 3D Plot with Deterministic Solution	
	Bibilography	20

#### 1 Introduction

This is a supplementary document for the report "Spatial epidemics dynamics: Synchronization" created for reproducibility. This supplement was created using knitr, an package that merges output languages such as LATEX with coding languages, allowing for dynamic reproducible report generation [1]. Other packages used are deSolve, titzDevice, and adaptivetau.

```
library("deSolve")
library("tikzDevice")
library("adaptivetau")
```

Output is currently suppressed during compile in the interest of speed - some code chunks take hours to run - but code can still be run using the uncompiled document SupplementaryMaterial.Rnw.

#### 2 BIFURCATION DIAGRAM

The bifurcation diagram was created using XPPAUT following the script described in the supplementary by Krylova and Earn [2]. In the script, equations for the single patch model with sinusoidal forcing were passed to the argument. The program was then instructed to solve the ODE by evaluating the equations with given initial conditions and parameters ranging over the parameter  $\mathcal{R}_0$  from 0-30 using Poincaré maps with period one year.

```
## Equations:
  f(x,y)=mu-R0*(gamma+mu)*(1 + a*cos(2*pi*t))*x*y-mu*x
  g(x,y)=R0*(gamma+mu)*(1 + a*cos(2*pi*t))*x*y-(gamma+mu)*y
  y(t+1)=g(x,y)
  R0(t+1)=R0
  ## Auxiliary Output Variables
  aux R0
  ## Initial Conditions
  init x=0.058824, y=0.00067043, R0=0
  par a=0.1, mu=0.02, gamma=28.077
  ## auxiliary parameters to corresponding with
## deterministic/stochastic models
  ## Plotting in XPPAUT
  @ xp=R0, yp=y
a_{19} @ x lo = 0, x hi = 40, y hi = 0.008, y lo = -0.008
  ## Range over R0
  @ rangeover=R0, rangestep=5000
  @ rangelow=0,rangehigh=30,rangereset=no
  @ rangeoldic=yes, lt=0
  ## Storing variables
  @ total = 650, trans = 0
  @ maxstor=10000000
  @ meth=discrete
  @ output=bifurcation.dat
  done
```

bifurcation.ode

When run using XPPAUT, this code outputs a file containing the values of time, susceptibles, infecteds, and  $\mathcal{R}_0$  for all specified iterations of  $\mathcal{R}_0$ .

This file can be generated and read by  $\mathbb{Q}$  via the following console commands:

```
system("xppaut bifurcation.ode -silent")
bfd <- read.table("bifurcation.dat", col.names=c("time", "S", "I", "RO"))</pre>
```

The bifurcation diagram can be generated using the following command:

```
plot(I$bfd,R0$bfd)
```

# 3 DETERMINISTIC SOLUTION

The SIR meta-patch vector field function, SIRmeta.vector.field, returns the values of  $\frac{dS_i}{dt}$ ,  $\frac{dI_i}{dt}$ , and  $\frac{dR_i}{dt}$  for all patches (i.e.  $1 \le i \le n$ ) at a given time, given the values of  $S_i$ ,  $I_i$ , and  $R_i$  for all patches and the values of parameters  $\mathcal{R}_0$  (the basic reproductive number),  $\mu$  (the death/birth rate),  $\gamma$  (the recovery rate),  $\alpha$  (the strength of seasonal forcing), m (the mixing parameter) and EC (a switch: if TRUE use equal coupling, otherwise use nearest neighbour coupling).

It creates the beta matrix:

$$\beta(t) = \langle \beta \rangle (1 + \alpha \cos(2\pi t)) M \tag{1}$$

where M is the mixing matrix. If equal coupling is used then

$$M = \begin{bmatrix} 1 - m & \frac{m}{n-1} & \frac{m}{n-1} & \frac{m}{n-1} \\ \frac{m}{n-1} & 1 & \frac{m}{n-1} & \frac{m}{n-1} \\ \frac{m}{n-1} & \frac{m}{n-1} & 1 \\ \vdots \\ \frac{m}{n-1} & & \ddots \end{bmatrix}$$

Otherwise nearest neighbour coupling is used.

$$M = \begin{bmatrix} 1 - m & \frac{m}{2} & 0 & 0 & \dots & \frac{m}{2} \\ \frac{m}{2} & 1 - m & \frac{m}{2} & 0 & & \vdots \\ 0 & \frac{m}{2} & 1 - m & & & \\ 0 & 0 & & \ddots & & \\ \vdots & & & \ddots & \frac{m}{2} \\ \frac{m}{2} & & & \dots & \frac{m}{2} & 1 - m \end{bmatrix}$$

The proportional dynamics of a single patch in the meta-patch SIR model are given by

$$\frac{dS_i}{dt} = \mu - S_i \sum_{j=1}^n \beta_{ij}(t) I_j - \mu S_i$$

$$\frac{dI_i}{dt} = S_i \sum_{j=1}^n \beta_{ij}(t) I_j - \gamma I_i - \mu I_i$$

$$\frac{dR_i}{dt} = \gamma I_i - \mu R_i$$
(2)

```
SIRmeta.vector.field <- function(t,vars,parms=NULL) {</pre>
  ##parms should be of form (R_0,gamma,mu,alpha,m,EC)
  ##vars should be of form (S,I,R)
  with(as.list(c(parms,vars)), { #So that it can call variables stored in parms
    ##vars contains the values of S, I, and R at time t
    n <- length(vars)/3</pre>
    S <- vars[1:n]
    I \leftarrow vars[(n+1):(2*n)]
    R \leftarrow vars[(2*n+1):(3*n)]
    #Equal Coupling
    if (EC==TRUE) {
      ##mixing matrix
      M \leftarrow matrix(m/(n-1),n,n)+(1-m*(1+1/(n-1)))*diag(n)
       ##beta matrix
      betam <- R_0*(gamma+mu)*(1+alpha*cos(2*pi*t))*M
    }else{
      #Nearest Neighbors
      ##mixing matrix
      M \leftarrow (1-m)*diag(n)
      M[row(M)\%n==(col(M)+1)\%n] < -m/2
      M[row(M)\%n==(col(M)-1)\%n] <- m/2
      ##beta matrix
      betam <- R_0*(gamma+mu)*(1+alpha*cos(2*pi*t))*M
```

```
dS <- NULL
dI <- NULL
dR <- NULL
for (i in 1:n){
    dS[i] <- mu*(1-S[i]) - S[i]*sum(betam[i,]*I) ##dS_i/dt
    dI[i] <- S[i]*sum(betam[i,]*I) - (mu+gamma)*I[i] ## dI_i/dt
    dR[i] <- gamma*I[i] -mu*R[i]
}
vec.fld <- c(dS=dS,dI=dI,dR=dR)
return(list(vec.fld))
})</pre>
```

The run.soln function returns the values of  $S_i$ ,  $I_i$ , and  $R_i$  for all patches at times given in times, given the initial conditions, the vector field function, and parameter to pass to the vector field function. It also returns the value of "coherence" at all times. A state is said to be coherent at time step t when  $I_1(t) = I_2(t) = ... = I_n(t)$  for all n patches [3]. The measure used here of "how coherent" a state is is the coefficient of variation, also called the relative standard deviation, of  $\{I_i(t)\}_{i=1}^n$ . It is the standard deviation of these values divided by the mean. The smaller it is the closer all states are to the mean and the more coherent the state is.

$$c_v(t) = \frac{SD(I_i(t))}{\overline{I_i(t)}}$$

$$= \frac{\sqrt{\frac{1}{n} \sum_{i=1}^{n} \left( I_i(t) - \overline{I_i(t)} \right)}}{\overline{I_i(t)}}$$
(3)

This is calculated using **@**'s native sd and mean functions.

Function plot.SIRmeta uses run.soln to get the solution of the determinitic model with the given initial conditions and parameters, and plots the results. It should be noted that when outputting the images, they were run through the tikzDevice package for to enable the use of LATEX within figures. The images were also saved to PDF through tikzDevice.

```
#Given ics and paramters, plot the det soln of the SIR meta model to pdf
plot.SIRmeta <- function(n,ic=c(S=rep(0,n),I=rep(0,n),R=rep(1,n)),tmax=1,
                      parms,...){
  ##parms should be of form (R_0,gamma,mu,alpha,m,EC)
 with(as.list(c(parms)), { #So that it can call variables stored in parms
  ## get solutions
 run <- run.soln(n=n,ic=ic,tmax=tmax,</pre>
           func=SIRmeta.vector.field,
           parms=parms)
  times<-run$times
  soln<-run$soln
  coherence<-run$coherence
  #Print to pdf using tikz
  filename <- paste0("images/det",if(EC) "EC" else "NN", "RO", R_O, "m", m, ".tex")
  if(file.exists(filename)){ #If the filename is already used
   filename <- pasteO("images/det",if(EC) "EC" else "NN", "RO", R_O, "m", m, "(1).tex")
  if(file.exists(filename)){ #If the filename(1) is already used
   filename <- paste0("images/det",if(EC) "EC" else "NN", "RO", R_O, "m", m, "(2).tex")
  tikz(filename, standAlone=TRUE, width=6, height=6)
  #leave room on right margin for coherence label
  par(mar = c(5,5,2,5))
  #Calculates I* to estimate how high the y-axis needs to be
  IEE \leftarrow (1 - 1/R_0)*mu/gamma
  ## draw box for plot
  plot(0,0,xlim=c(0,tmax),ylim=c(0,IEE*5),type="n",xlab="Time (years)",
    ylab="Prevalence (I)",las=1)
  #Plot I for all patches with different colours and lty
  for (i in 1:n){
   lines(times,soln[,pasteO('I',i)],col=i,lty=i)
  #Plot coherence over top
  par(new = T)
  plot(times,coherence,type='1',lwd=2,axes=F,xlab=NA, ylab=NA,ylim=c(0,1))
  axis(side=4)
  #Label left axis
  mtext(side = 4, line = 3, 'Coherence')
  #Make legend
 labels \leftarrow c(1:n)
  for(i in 1:n){
   labels[i] <-paste('$I_{',i,'}$',collapse=NULL)</pre>
  legend("topright",legend=labels,col=1:n,lty=1,bty='n')
  tools::texi2dvi(filename,pdf=T)
  })
```

}

The following chunk of code uses plot.SIRmeta to plot solutions to the meta-patch model for different parameters and with initial conditions within  $\pm 30\%$  of the Endemic Equilibrium (EE) to preserve asymptotic local stability of the system[4]. This is so that the solutions converge quickly to a stable solution. The endemic equilibrium is

$$S^* = \frac{1}{\mathcal{R}_0}$$

$$I^* = \frac{\mu}{\gamma} \left( 1 - \frac{1}{\mathcal{R}_0} \right)$$

$$R^* = 1 - S^* - I^*$$

$$(4)$$

```
#Parameters
R_0 < -17
           ##basic reproduction rate (if the system had no forcing)
gamma <- 365/13 ##inverse of mean infectious period
mu <- 1/50 ##death and birth rate
alpha <- 0.1 ##strength of seasonal forcing
n <- 10 ## Number of patches
m <- 0.2 ##Connectivity matrix parameter
EC <- FALSE ##if true use equal coupling, if false use nearest neighbors
##Initial conditions
#Want ICs near EE for stability. Find EE values:
SEE <-1/R_0
IEE <-(1 - 1/R_0)*mu/gamma
REE <-1-SEE-IEE
#Use random initials conditions within +/-30% of EE values
set.seed(34) #For reproducbility
percent <- 0.3
SO <- runif(SEE*(1-percent), SEE*(1+percent), n)
IO <- runif(IEE*(1-percent), IEE*(1+percent), n)</pre>
RO <- 1-SO-IO
tmax<-15
#### Plotting lots of solutions to pdf ####
#Making detNNR017m0.2
plot.SIRmeta(n=n,ic=c(S=S0,I=I0,R=R0),tmax=tmax,
                      parms=c(R_0=R_0,gamma=gamma,mu=mu,alpha=alpha,m=m,EC=EC))
#Making detNNR010m0.2
R 0<-10
plot.SIRmeta(n=n,ic=c(S=S0,I=I0,R=R0),tmax=tmax,
                      parms=c(R_0=R_0,gamma=gamma,mu=mu,alpha=alpha,m=m,EC=EC))
#Making detNNR025m0.2
R_0<-25
plot.SIRmeta(n=n,ic=c(S=S0,I=I0,R=R0),tmax=tmax,
                      parms=c(R_0=R_0,gamma=gamma,mu=mu,alpha=alpha,m=m,EC=EC))
#Making detECR017m0.2s
R_0<-17
tmax<-5
EC<-TRUE
plot.SIRmeta(n=n,ic=c(S=S0,I=I0,R=R0),tmax=tmax,
```

```
parms=c(R_0=R_0,gamma=gamma,mu=mu,alpha=alpha,m=m,EC=EC))
#Making detNNR017m0.2s
EC<-FALSE
plot.SIRmeta(n=n,ic=c(S=S0,I=I0,R=R0),tmax=tmax,
                      parms=c(R_0=R_0,gamma=gamma,mu=mu,alpha=alpha,m=m,EC=EC))
#Making detNNR017m0.01
tmax<-15
m < -0.01
plot.SIRmeta(n=n,ic=c(S=S0,I=I0,R=R0),tmax=tmax,
                      parms=c(R_0=R_0,gamma=gamma,mu=mu,alpha=alpha,m=m,EC=EC))
#Making detNNR017m0.01wider
percent <- 0.4
SO <- runif(SEE*(1-percent), SEE*(1+percent), n)
IO <- runif(IEE*(1-percent), IEE*(1+percent), n)
RO <- 1-SO-IO
plot.SIRmeta(n=n,ic=c(S=S0,I=I0,R=R0),tmax=tmax,
                      parms=c(R_0=R_0,gamma=gamma,mu=mu,alpha=alpha,m=m,EC=EC))
```

#### 4 STOCHASTIC SIMULATION

Stochastic simulations of the model were also used.

### 4.1 Gillespie Algorthim

The Gillespie algorithm is a method of exactly simulating a stochastic process. Consider a process involving transitions between different states. The rate for a given transition i is labelled  $a_i$ . The total transistion rate is defined as

$$a_0 = \sum a_i \tag{5}$$

If we assume that the time spent in a given state is exponentially distributed, then by sampling from an exponential distribution with rate  $a_0$  (using  $\mathbb{Q}$ 's rexp function) we can simulate how much time was spent in a given state before the next transition.

The probability that transition i occured is  $\frac{a_i}{a_0}$ . By sampling from a uniform distribution over  $[0, a_0]$  we can simulate which transition occured in the following way: if the sampled value is in the  $i^{th}$  interval from the list

$$[0, a_1), [a_1, a_1 + a_2), \dots [a_1 + a_2 + \dots + a_{i-1}, a_1 + a_2 + \dots + a_i), \dots$$
 (6)

than say transition i occured.

For this problem, the states are the values of  $\{(S_i, I_i, R_i)\}_{i=1}^n$ . The state space is all possible set of n triplets of non-negative integers. Possible transitions are:

- A birth  $(S_i \to S_i + 1)$
- A transmission  $(S_i \to S_i 1 \text{ and } I_i \to I_i + 1)$
- A recovery  $(I_i \to I_i 1 \text{ and } R_i \to R_i + 1)$
- A death of a susceptible  $(S_i \to S_i 1)$
- A death of an infected  $(I_i \to I_i 1)$
- A death of a recovered  $(R_i \to R_i 1)$

for  $1 \le i \le n$ . The transition rates are:

• The birth rate  $\mu(S_i + I_i + R_i)$ 

- The transmission rate  $\frac{S_i}{S_i + I_i + R_i} \sum_{j=1}^n \beta_{ij}(t) I_j$
- The recovery rate  $\gamma I_i$
- The susceptible death rate  $\mu S_i$
- The infected death rate  $\mu I_i$
- The recovered death rate  $\mu R_i$

#### for $1 \leq i \leq n$ .

The function SIRmeta.Gillespie implements a stochastic simulation of the meta-patch SIR model using the Gillespie algorithm. It returns a list of times (starting at zero and ending near the input variable tmax with non-uniform step sizes) and the values of  $S_i$ ,  $I_i$ , and  $R_i$  for all patches at these times given initial conditions and parameter values.

```
SIRmeta.Gillespie <- function(tmax,ic=c(S0,I0,R0),</pre>
                                parms=c(R_0=2,gamma=0.25,mu=4e-5,
                                         alpha=0.1,m=0.2,EC=TRUE)){
  times <-0
  n \leftarrow length(ic)/3
  #S, I, R matrices: column index is patch index and rows (added later in while loop)
  #are values at different time steps
  S <- matrix(0,1,n)</pre>
  I \leftarrow matrix(0,1,n)
  R <- matrix(0,1,n)</pre>
  #Initial Coniditions
  S[1,] \leftarrow ic[1:n]
  I[1,] \leftarrow ic[(n+1):(2*n)]
  R[1,] \leftarrow ic[(2*n+1):(3*n)]
  index <- 1
  while(times[index] < tmax){</pre>
    #A matrix of all rates
    rates <- matrix(0,n,6)
    #the first column is all birth rates
    rates[,1] <- mu*(S[index,]+I[index,]+R[index,])</pre>
    #thrid column are recovery rates
    rates[,3] <- gamma*I[index,]</pre>
    #last three columns are death rates from S and I
    rates[,4] <- mu*S[index,]</pre>
    rates[,5] <- mu*I[index,]
    rates[,6] <- mu*R[index,]
    #Second column are transmission rates
    #Equal Coupling
    if(EC){
      M \leftarrow matrix(m/(n-1),n,n)+(1-m*(1+1/(n-1)))*diag(n) ##connectivity matrix
      betam <- R_0*(gamma+mu)*(1+alpha*cos(2*pi*times[index]))*M ##beta matrix
    }else{
      #Nearest Neighbors
      M \leftarrow (1-m)*diag(n)
      M[row(M)\%n==(col(M)+1)\%n] <- m/2
      M[row(M)\%n==(col(M)-1)\%n] <- m/2
      betam <- R_0*(gamma+mu)*(1+alpha*cos(2*pi*times[index]))*M ##beta matrix
```

```
rates[,2] <- S[index,]*colSums(t(betam[1:n,])*I[index,])/(S[index,]+I[index,]+R[index,])</pre>
totalrate <- sum(rates)</pre>
#Getting a timestep using R's exp distribution
timestep <- rexp(n=1, rate= totalrate)</pre>
times <- rbind(times, times[index]+timestep)</pre>
#Adding new row to S, I, R
S <- rbind(S, S[index,])
I <- rbind(I, I[index,])</pre>
R <- rbind(R, R[index,])</pre>
#Getting which compartment transistion occurred
#TO DO: Do a bisection search (or some other faster method) instead
#Get a random number from a uniform distribution
randomvar <- runif(n=1,min=0,max=totalrate)</pre>
interval <- 1
#Make the rates matrix into a vector for conveince
rateslisted <- c(0,as.vector(t(rates)))</pre>
#Find which interval randomval is in from list:
\#[0,a_1), [a_1, a_1+a_2), etc
#If in interval [a_1 + ... + a_i (i-1), a_1 + ... + a_i),
#transition given by rate a_i occurred
while(interval < (6*n)){</pre>
  if(randomvar >= sum(rateslisted[1:interval]) &
     randomvar < sum(rateslisted[1:(interval+1)])){</pre>
    break
  }else{
    interval<- interval+1
#Get row, col indices of a_i in rates matrix
#row index says in which patch a transition occurred
#col index says what type of transition occurred
colInd <- interval%%6</pre>
rowInd <- ceiling(interval/6)</pre>
if(colInd ==1){ ##A birth occurred
  S[index+1,rowInd] <- S[index,rowInd]+1
}else if(colInd ==2){ ##An infection occurred
  S[index+1,rowInd] <- S[index,rowInd]-1
  I[index+1,rowInd] <- I[index,rowInd]+1</pre>
}else if(colInd ==3){ ##A recovery occurred
  I[index+1,rowInd] <- I[index,rowInd]-1</pre>
  R[index+1,rowInd] <- R[index,rowInd]+1</pre>
}else if(colInd ==4){ ##A death in S occurred
  S[index+1,rowInd] <- S[index,rowInd]-1</pre>
}else if(colInd ==5){ ##A death in I occurred
  I[index+1,rowInd] <- I[index,rowInd]-1</pre>
}else if(colInd ==0){ ##A death in R occurred
  R[index+1,rowInd] <- R[index,rowInd]-1</pre>
```

```
index <- index+1
}
return(cbind(times, S, I, R))
}</pre>
```

The following code uses the SIRmeta. Gillespie function to plot a stochastic simulation of the model.

```
#For reprodubility set the seed
set.seed(899)
tmax <- 10
n < -10
#Parameters
R_0 \leftarrow 17 ##basic reproduction rate (if the system had no forcing)
gamma <- 365/13 ##inverse of mean infectious period
mu <- 1/50 ##death and birth rate
alpha <- 0.1 ##strength of seasonal forcing
n <- 10 ## Number of patches
m <- 0.5 ##Connectivity matrix parameter
EC <- TRUE ##if true use equal coupling, if false use nearest neighbors
##Initial conditions
#Want ICs near EE for stability. Find EE values:
SEE <-1/R_0
IEE <-(1 - 1/R_0)*mu/gamma
REE <-1-SEE-IEE
#Sample from initials conditions within +/-10% of EE values
#convert from proportions to numbers and convert to integers
percent <- 0.1
population <- 3000
possibleS0 <- seq(as.integer(population*SEE*(1-percent)),</pre>
                  as.integer(population*SEE*(1+percent)),1)
possibleI0 <- seq(as.integer(population*IEE*(1-percent)),</pre>
                  as.integer(population*IEE*(1+percent)),1)
SO <- sample(possibleSO, n,replace=TRUE)
IO <- sample(possibleIO, n,replace=TRUE)</pre>
RO <- population-SO-IO
#Get simulation results
results <- SIRmeta.Gillespie(tmax,ic=c(S0,I0,R0),
                              parms=c(R_0,gamma,mu,alpha,m,EC))
times<-results[,1]</pre>
I \leftarrow results[,(n+2):(2*n+2)]
#Save data for later use and plot to pdf and png
write.table(results,file="GillECR017m0.2.pdf")
pdf(file="images/GillECR017m0.2.pdf")
copy <- dev.cur()</pre>
png(file="images/GillECR017m0.2.png")
dev.control("enable")
## draw box for plot
 plot(0,0,xlim=c(0,tmax),ylim=c(0,population*IEE*2),xlab="Time (years)",
```

```
ylab="Prevalence (I)")
#Plot all I_i using a differnt line colour and lty for each patch
for (i in 1:n) {
    lines(times, I[,i],col=i,lwd=0.2)
}
#Make legend
legend("topright",legend=1:n,fill="white",title="Patch",col=1:n,lty=1,cex=0.5)

dev.copy(which=copy)
dev.off
dev.off
```

This code runs very slowly for large populations. For a 250,000 person per patch population it took 3 hours. This method for simulating the stochastic realization of the model is thus not practial.

# 4.2 Adaptive Tau-Leaping Algorihim

To simulate a stochastic process efficiently, an approximate method can be used. The adaptive tau-leaping algorithm works by

The runadaptau function contains a list of all possible transitions and rateFct, a function that returns the values of all transition rates given the time and current state. runadaptau passes these, along with the input initial conditions and parameters, to the ssa.adaptivetau function. It returns a list of times (starting at zero and ending near input tmax with non-uniform step size) and the values of the state at all these times.

```
run.adaptau <-function(tmax,ic=c(S0,I0,R0),
                            params=c(R_0,gamma,mu,alpha,m,EC)){
#Get the # of patches from the ics
n \leftarrow length(ic)/3
#TO DO: This should be changed so that it works with different n values
#and is less ugly
#A list of all possible transitions
transitions <- list(c(S1 = +1), c(S1 = -1, I1 = +1), c(I1 = -1, R1 = +1), c(S1 = -1),
                    c(I1 = -1), c(R1 = -1),
                  c(S2 = +1), c(S2 = -1, I2 = +1), c(I2 = -1, R2 = +1), c(S2 = -1),
                  c(I2 = -1), c(R2 = -1),
                  c(S3 = +1), c(S3 = -1, I3 = +1), c(I3 = -1, R3 = +1), c(S3 = -1),
                  c(I3 = -1), c(R3 = -1),
                  c(S4 = +1), c(S4 = -1, I4= +1), c(I4 = -1, R4= +1), c(S4 = -1),
                  c(I4 = -1), c(R4 = -1),
                  c(S5 = +1), c(S5 = -1, I5 = +1), c(I5 = -1, R5 = +1), c(S5 = -1),
                  c(I5 = -1), c(R5 = -1),
                  c(S6 = +1), c(S6 = -1, I6 = +1), c(I6 = -1, R6 = +1), c(S6 = -1),
                  c(16 = -1), c(R6 = -1),
                  c(S7 = +1), c(S7 = -1, I7 = +1), c(I7 = -1, R7 = +1), c(S7 = -1),
                  c(I7 = -1), c(R7 = -1),
                  c(S8 = +1), c(S8 = -1, I8 = +1), c(I8 = -1, R8 = +1), c(S8 = -1),
                  c(I8 = -1), c(R8 = -1),
                  c(S9 = +1), c(S9 = -1, I9 = +1), c(I9 = -1, R9 = +1), c(S9 = -1),
                  c(I9 = -1), c(R9 = -1),
                  c(S10 = +1), c(S10 = -1, I10 = +1), c(I10 = -1, R10 = +1), c(S10 = -1),
                  c(I10 = -1), c(R10 = -1))
```

```
#A function that outputs the transition rates given parameters, S,I,R, and t
rateFct <- function(x,params,time){</pre>
  #x should be of form: x=c(S,I,R)
  #parms should be of form: paarms=c(R_0,gamma,mu,alpha,m,EC)
  with(
     as.list(c(params)),
  n \leftarrow length(x)/3
  S \leftarrow as.vector(x[1:n])
  I \leftarrow as.vector(x[(n+1):(2*n)])
  R \leftarrow as.vector(x[(2*n+1):(3*n)])
  #A matrix of all rates
  rates <- matrix(0,n,6)
  #the first column is all birth rates
  rates[,1] <- mu*(S+I+R)
  #thrid column are recovery rates
  rates[,3] <- gamma*I</pre>
  #last three columns are death rates from S and I
  rates[,4] <- mu*S
  rates[,5] <- mu*I
  rates[,6] <- mu*R
  #Second column are transmission rates
  #Equal Coupling
  if (EC==TRUE) {
     \texttt{M} \leftarrow \mathtt{matrix}(\texttt{m}/(\texttt{n-1}),\texttt{n},\texttt{n}) + (1-\texttt{m}*(1+1/(\texttt{n-1})))*\mathtt{diag}(\texttt{n}) \quad \#\texttt{connectivity matrix} 
    betam <- R_0*(gamma+mu)*(1+alpha*cos(2*pi*time))*M ##beta matrix
  }else{
    #Nearest Neighbors
    M \leftarrow (1-m)*diag(n)
    M[row(M)\%n==(col(M)+1)\%n] <- m/2
    M[row(M)\%n==(col(M)-1)\%n] <- m/2
    betam <- R_0*(gamma+mu)*(1+alpha*cos(2*pi*time))*M ##beta matrix
  rates[,2] \leftarrow S*colSums(t(betam[1:n,])*I)/(S+I+R)
  return(as.vector(t(rates)))
init.values <- c(S=S0,I=I0,R=R0)</pre>
#Get approximate simulation
simResults <- ssa.adaptivetau(init.values,transitions, rateFct, params, tf=tmax)</pre>
I <- simResults[,(12:21)]</pre>
  #Measure of coherence: the coefficient of variation/relative
#standard deviation of all the patches at each time step
coherence <- rep(0,length(simResults[,1]))</pre>
for(tims in 1:length(simResults[,1])){
  coherence[tims] <- sd(I[tims,])/mean(I[tims,])</pre>
  #In case the mean is zero (and thus the coefficient of variation becomes nan or inf)
```

```
if(is.nan(coherence[tims]) | is.infinite(coherence[tims])){
    coherence[tims]<- 0</pre>
#Is the disease extinct (I pop below 20% of IEE for awhile) in any of the patches?
#Store 1s for yes, 0 for nos
localextinction<-c(1:n)</pre>
for(i in 1:ncol(I)){
  if(mean(I[(nrow(I)-50):nrow(I),i]) <= IEE*population*0.2){</pre>
    localextinction[i]<-1</pre>
  }else{
    localextinction[i] <-0</pre>
localext<-mean(localextinction)</pre>
#If the disease is extinct in all patches there is global extinction
if(all(localextinction==1)){
    globalextinction<-1</pre>
  }else{
    globalextinction<-0</pre>
return(list("sim" = simResults, "coherence"= coherence, "localext"=localext,
             "globalext"=globalextinction))
```

The following function,plot.adaptau, uses runadaptau to approximately simulate the meta-patch SIR model and plot the results.

```
plot.adaptau <- function(tmax,init.values,params,IEE,population){</pre>
 with(as.list(c(parms)), {
  run <- run.adaptau(tmax,init.values,params)</pre>
  simResults <- run$sim
  coherence <- run$coherence
  ##Saving the data and the print plot to pdf
  filename <- paste0("adaptau",if(EC==TRUE) "EC" else "NN", "RO", R_O, "m", m)
  if(file.exists(filename)){ #If the filename is already used
   filename <- paste0("adaptau",if(EC==TRUE) "EC" else "NN", "RO", R_O, "m", m, "(1)")
  if(file.exists(filename)){ #If the filename(1) is already used
   filename <- pasteO("images/adaptau",if(EC==TRUE) "EC" else "NN", "RO", R_O, "m", m,
                       "(2)")
  write.table(run,file=paste0(filename,".csv"))
  pdf(file=paste0("images/",filename,".pdf"))
  ## draw box for plot
  #leave room on right margin for Coherence label
  par(mar = c(5,5,2,5))
```

```
#Solutions should ocsillate around IEE so ylim should go to around twice
#that so solns are in the middle of plot
plot(0,0,xlim=c(0,tmax),ylim=c(0,population*IEE*4),xlab="Time (years)",
     ylab="Prevalence (I)")
##plot I_i for each patch
j<-1:n
for (i in 1:n) {
 if(i\%8==1) j[i] <- i+3 #indices so that black and solid lines are
 #skipped; saved for legend
 lines(simResults[,1], simResults[,11+i],col=j[i],lty=j[i],lwd=0.5)
 # use a different line colour and line type for each patch
#Currently unused code for plotting the det soln over this sim
 # draw.single.soln(n,ic=c(S=SEE,I=IEE,R=(1-SEE-IEE)),tmax=tmax,
           func=SIR.vector.field,
           parms=c(R_0,gamma,mu,alpha,population))
#Plot coherence onto of pre-existing plot
par(new = T)
plot(simResults[,1],coherence,type='l',axes=F,xlab=NA, ylab=NA, ylim=c(0,1))
#y axis and coherence label on right axis
axis(side=4)
mtext(side = 4, line = 3, 'Coherence')
#Make legends
legend("topright",legend=1:n,title="Patch",bg='white',col=j,lty=j,lwd=0.5,cex=0.8)
legend("bottomright", legend="Relative Standard Deviation",bg='white',
     col=1,lty=1, cex=0.8)
dev.off()
})
```

#### Using plot.adaptau to simulate the model with different parameter values:

```
#Parameters
R_0 \leftarrow 17 ##basic reproduction rate (if the system had no forcing)
gamma <- 365/13 ##inverse of mean infectious period
mu <- 1/50 ##death and birth rate
alpha <- 0.1 ##strength of seasonal forcing
n <- 10 ## Number of patches
m <- 0.2 ##Connectivity matrix parameter
EC <- TRUE ##if true use equal coupling, if false use nearest neighbors
params<-c(R_0,gamma,mu,alpha,m,EC)</pre>
##Initial conditions
#Want ICs near EE for stability. Find EE values:
SEE <-1/R_0
IEE <-(1 - 1/R_0)*mu/gamma
REE <-1-SEE-IEE
 #Sample ics from points +/-10% from the EE values
percent <- 0.1
```

#### 5 COHERENCE DEPENDENCE ON PARAMETERS

The dependence of a meta-patch solution becoming "coherent" on the parameters used in the model was explored. A system is considered coherent when the mean value of  $c_v(t)$  from t = 9 years to t = 10 years is below a threshold value. We will refer to this value as the coherence threshold.

A list of  $\mathcal{R}_0$  (the basic reproductive number) values from 1.1 to 30 was made, along with a list of m (the mixing parameter) values. For each set of  $\mathcal{R}_0$  and m values, a solution or simulation of the model was run ten times each with different initial conditions (all within 10% of the Endemic Equilibrium values). It was stored whether or not each solution or simulation became coherent; a one was saved if it did and a zero was saved otherwise. By taking the mean of these values for all runs, the probability of becoming coherent for a given  $\mathcal{R}_0$  and m value was found.

If parameter DetOrAdap is TRUE this is done using deterministic solutions to the model, created by the run.soln function. Otherwise, this is done using adaptive tau simulations in the runadaptau function.

```
coherenceGrid <- function(params=c(gamma,mu,alpha,EC),</pre>
                              R_Olist,mlist,nsample,DetOrAdap,population=1){
  with(
       as.list(c(params)),
  n <- 10 ## Number of patches
  ##Param for Initial conditions
  percent <- 0.1
  tmax <- 10
  gridsizem<-length(mlist)</pre>
  gridsizeR_0 <- length(R_0list)</pre>
  avgCoherence<- matrix(0, gridsizem,gridsizeR_0)</pre>
  avgLE<- matrix(0, gridsizem,gridsizeR_0)</pre>
  avgGE<- matrix(0, gridsizem,gridsizeR_0)</pre>
  index2 <- 1
  for (R_0 in R_0list){
    index1 <- 1
     #Want ICs near EE for stability. Find EE values:
     #Note: They depend on R_O
     SEE <-1/R 0
     IEE <-(1 - 1/R_0)*mu/gamma
```

```
if(!DetOrAdap){ #Only need these for adaptau sim
      possibleS0 <- seq(as.integer(population*SEE*(1-percent)),</pre>
                          as.integer(population*SEE*(1+percent)),1)
      possibleI0 <- seq(as.integer(population*IEE*(1-percent)),</pre>
                          as.integer(population*IEE*(1+percent)),1)
    for (m in mlist){
      coherenceProb <- rep(0, nsample)</pre>
      coherenceProb <- rep(0, nsample)</pre>
      localextProb <- rep(0, nsample)</pre>
      globalextProb <- rep(0, nsample)</pre>
      for (sampler in 1:nsample){
       #Get det soln nsample times with different ICs
       #Take mean of coherence near end of soln
       #(since coherence can be somewhat oscilatory)
        if(DetOrAdap){
           SO <- runif(n,SEE*(1-percent),SEE*(1+percent))
           IO <- runif(n, IEE*(1-percent), IEE*(1+percent))</pre>
           RO <- 1-SO-IO
           coherenceRun <- run.soln(n,ic=c(S=S0,I=I0,R=R0),tmax=tmax,
                             func=SIRmeta.vector.field,
                           parms=c(R_0,gamma,mu,alpha,m,EC))$coherence
        }else{
          SO <- sample(possibleSO, n,replace=TRUE)
          IO <- sample(possibleIO, n,replace=TRUE)</pre>
          RO <- population-SO-IO
          run <- run.adaptau(tmax,init.values=c(S=S0,I=I0,R=R0),</pre>
                                      params=c(R_0,gamma,mu,alpha,m,EC))
          coherenceRun<-run$coherence
          localextProb[sampler] <- run$localext</pre>
          globalextProb[sampler] <- run$globalext</pre>
      mean <- mean(coherenceRun[length(coherenceRun)-(100:0)])</pre>
      if (mean<threshold) {</pre>
          coherenceProb[sampler] <- 1</pre>
          coherenceProb[sampler] <- 0</pre>
   #Take mean of all coherence for all runs with these parameters
   avgCoherence[index1,index2] <- mean(coherenceProb)</pre>
   avgLE[index1,index2] <- mean(localextProb)</pre>
   avgGE[index1,index2] <- mean(globalextProb)</pre>
   index1<-index1+1
 index2<-index2+1
#Save data
filename<-paste0("grid",if(DetOrAdap) "det" else "adap",if(EC) "EC" else "NN",".csv")
```

#### 5.1 3D Plot with Deterministic Solution

A 3D plot showing the probability of coherence vs  $\mathcal{R}_0$  and m is created in the function coherence 3DPlot.

```
coherence3DPlot <- function(avgCoherence,params=c(gamma,mu,alpha,EC),R_0list,</pre>
                            mlist,nsample,DetOrAdap,population){
  with(
       as.list(c(params)),
  #3D plot:
  filename < -paste0("images/",if(DetOrAdap) "det" else "adap","3D",if(EC) "EC" else "NN",
                   ".pdf")
  pdf(file=filename)
  #Making a colour gradient for the plot
  nbcol <- 100
  #You can change start and end to get different parts of the rainbow
  color <- rev(rainbow(nbcol, start = 3/6, end = 3.5/6))</pre>
  #The colour gradient will correspond to the 'height' of the plot
 zcol <- cut(t(avgCoherence), nbcol)</pre>
 persp( R_Olist,mlist, t(avgCoherence), theta = 30, phi = 30, expand = 0.5,
        col = color[zcol], ltheta = 120, shade = 0.75, ticktype = "detailed",
      xlab = "Basic Reproduction Number", ylab = "Mixing Parameter",
     zlab = "Probability of Coherence")
  dev.off()
       })
```

The following code calls coherence3DPlot to make a plot.

```
#Use the det soln if TRUE or adaptau sim if FALSE

DetOrAdap <- TRUE

#The value at which things are considered coherent
threshold <- 0.05
#threshold <- 0.15 #for adaptau higher tol needed

#Parameter values for grid
gridsizeR_0 <- 100
gridsizem <- 100
nsample <- 10 #Number of runs per grid pt
mlist <- seq(0.0001, 0.15, (0.15-0)/(gridsizem-1))
R_Olist <- seq(1.1,30,(30-1.1)/(gridsizeR_0-1))
```

```
#Parameters
gamma <- 365/13 ##inverse of mean infectious period
mu <- 1/50 ##death and birth rate
alpha <- 0.1 ##strength of seasonal forcing
EC <- FALSE ##if true use equal coupling, if false use nearest neighbors
#Get data
data <- coherenceGrid(params=c(gamma=gamma,mu=mu,alpha=alpha,EC=EC),</pre>
                      R_Olist, mlist, nsample, DetOrAdap, population)
avgCoherence <- data$avgCoherence</pre>
#Plot
coherence3DPlot(avgCoherence,params=c(gamma=gamma,mu=mu,alpha=alpha,EC=EC),
                R_Olist,mlist,nsample,DetOrAdap,population=1)
data <- coherenceGrid(params=c(gamma=gamma,mu=mu,alpha=alpha,EC=TRUE),</pre>
                      R_Olist, mlist, nsample, DetOrAdap, population)
avgCoherence <- data$avgCoherence</pre>
coherence3DPlot(avgCoherence,params=c(gamma=gamma,mu=mu,alpha=alpha,EC=TRUE),
                R_Olist,mlist,nsample,DetOrAdap,population=1)
```

#### 5.2 Four Panel Plot with Adaptive Tau

The function fourPanPlot makes a four panel plot showing the probability of coherence, the probability of local and global extinction, overtop of the bifurcation diagram of the single patch forced SIR model.

```
fourPanPlot <- function(avgCoherence,avgLE,avgGE,</pre>
                          params=c(gamma,mu,alpha,EC),R_Olist,mlist,
                          nsample,DetOrAdap,population){
   with(
       as.list(c(params)),
   if (length(mlist)!=4){
     print("Warning: mlist should have length 4 for 4 panel plot
       \\ The first 4 elements of mlist will be used")
#Save plot to pdf
filename<-paste0("images/",if(DetOrAdap) "det" else "adap","4pan",
                  if(EC) "EC" else "NN",".pdf")
   pdf(file=filename)
#Get bifur data for backgrounds
bfd <- read.table("bifurcation.dat",</pre>
                   col.names=c("time", "S", "I", "RO", "log10S", "log10I"))
#Get max probs for y lim
maxExt <- max(avgLE,avgGE)</pre>
maxCoh <- max(avgCoherence)</pre>
#To fit bifur data to probability plot
bfdI <- maxExt*(1.1/max(bfd$I))*bfd$I</pre>
```

```
#leave room on outside margins for labels on rigth, left and bottom
\#par(mar = c(5,5,2,5))
par(oma=c(4,4,2,4))
#A four panel plot
par(mfrow=c(2,2))
par(mar = c(2, 3, 1, 2))
par(tcl = -0.25)
for(ind in 1:4){
  #Make an empty plot
  plot(0,0,xlim=c(1,30),
              vlim=c(0,maxExt*1.1),xlab=NA, ylab=NA,type='n')
   #Plot the bifur background
  points(bfd$R0,bfdI, pch=20, col="grey90", xpd=FALSE)
   #Plot LE, GE probs
  lines(R_Olist, avgLE[ind,],col=4)
   lines(R_Olist, avgGE[ind,],col=2)
   #Plot coherence prob overtop of current panel
   par(mfg = c(ceiling(ind/2), (ind-1)\%2 +1))
  plot(R_Olist, avgCoherence[ind,],type='1',col=3,axes=F,xlab=NA,
       ylab=NA,ylim=c(0,maxCoh*1.1))
   axis(side=4)
   #Label m values on each plot
  mtext(pasteO("m=",mlist[ind]), side = 3, line = -2, adj = 0.1,
        cex = 0.8, col = 1)
#Axes labels:
mtext("Basic Reproduction Number", side = 1, outer = TRUE, line = 1)
#Left axis label has multiple colors
mtext(expression("Probability of "*phantom("Local/ Global Extinction")),
      side = 2, outer = TRUE, line = 1, col = 1)
mtext(expression(phantom("Probability of ")*"Local"*phantom("/ Global Extinction")),
      side = 2, outer = TRUE, line = 1, col = 4)
mtext(expression(phantom("Probability of Local")*"/"*phantom(" Global Extinction")),
      side = 2, outer = TRUE, line = 1, col = 1)
mtext(expression(phantom("Probability of Local/")*" Global"*phantom(" Extinction")),
      side = 2, outer = TRUE, line = 1, col = 2)
mtext(expression(phantom("Probability of Local/ Global")*" Extinction"),
      side = 2, outer = TRUE, line = 1, col = 1)
mtext("Probability of Coherence", side = 4, outer = TRUE, line = 1, col = 3)
 dev.off()
      })
```

#### Using fourPanPlot:

```
#The value at which things are considered coherent
threshold <- 0.15
#Parameter values for grid
gridsizeR_0 <- 200
```

```
gridsizem <- 4
nsample <- 20
mlist < c(0.2,0.45,0.7,0.9)
R_0list <- seq(1.1,30,(30-1.1)/(gridsizeR_0-1))
DetOrAdap <- FALSE</pre>
 #Parameters
gamma <- 365/13 ##inverse of mean infectious period
mu <- 1/50 ##death and birth rate
alpha <- 0.1 ##strength of seasonal forcing
n <- 10 ## Number of patches
EC <- FALSE ##if true use equal coupling, if false use nearest neighbors
population <- 250000
#Get data
data <- coherenceGrid(params=c(gamma=gamma,mu=mu,alpha=alpha,EC=EC),</pre>
                       R_Olist,mlist,nsample,DetOrAdap,population)
avgCoherence <- data$avgCoherence</pre>
avgLE <- data$avgLE</pre>
avgGE <- data$avgGE</pre>
   #Plot
fourPanPlot(avgCoherence,avgLE,avgGE,
            params=c(gamma=gamma,mu=mu,alpha=alpha,EC=EC),
                         R_Olist,mlist,nsample,DetOrAdap,population)
```

#### REFERENCES

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