

# Modeling the Effect of School Closures in a Pandemic Scenario: Exploring Two Different Contact Matrices

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## R codes

### Introduction

As supplementary materials to the article, this PDF file contains four R codes (.r files):

1. “Flu\_School\_Closure\_CID.r”: This is the main R code for ordinary differential equation model.
2. “Flu\_School\_Closure\_CID\_Data.r”: This is the R code that creates the summary result file.
3. “Flu\_School\_Closure\_CID\_Plot.r”: This is the R code that plots the figures.
4. “Flu\_School\_Closure\_CID\_Difference\_eqn.r”: This is the R code for the difference equation model.

Note: These codes are provided as examples for readers to use in their own studies. These sample codes are by no means the most efficient ways of writing R codes. There are likely to be more than one way to program in R to achieve the same goal and readers may re-write the codes in a more efficient manner (with fewer lines).

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## R code 1: “Flu\_School\_Closure\_CID.r”

```
#####  
## This model is written by Dr. Isaac Chun Hai Fung  
## With input from Dr. Manoj Gambhir and Dr. John Glasser on the mixing matrix  
##  
## The model is based on a previously published model:  
## Fung IC-H, Antia R, Handel A (2012)  
## How to Minimize the Attack Rate during Multiple Influenza Outbreaks in a  
## Heterogeneous Population.  
## PLoS ONE 7(6): e36573. doi:10.1371/journal.pone.0036573  
##  
## This R code was based on the R code previously written for Fung, Antia and Handel  
## (2012). The model is now expanded to have 4 age groups instead of 2.  
##  
## Final edits were made on November 21, 2014 for the purpose of publication.  
#####  
  
rm(list=ls())  
# This clears the workspace to make sure no leftover variables are  
# floating around. It is not strictly needed but it is often a good idea.  
  
graphics.off();  
# Close all graphics windows, in case there are still  
# some open from previous work that we did  
  
library(deSolve)  
# Load ODE solver package. You need to have the package installed first.  
  
#####  
## First, we specify the function that describes the differential equation model  
## for the simulated virus infection.  
##  
## This function is called by lsoda (the ODE solver) in the main program  
#####  
  
odeequations=function(t,y,parameters)  
## The function has to be written in a certain form, dictated by lsoda  
{  
  S1=y[1]; S2=y[2]; S3=y[3]; S4=y[4]; # Susceptible  
  I1=y[5]; I2=y[6]; I3=y[7]; I4=y[8]; # Infected (Infectious)  
  R1=y[9]; R2=y[10]; R3=y[11]; R4=y[12]; # Recovered  
  
  ## Model parameters, passed into function by main program  
  beta11 = parameters[1];  
  beta12 = parameters[2];  
  beta13 = parameters[3];  
  beta14 = parameters[4];  
  beta21 = parameters[5];  
  beta22 = parameters[6];  
  beta23 = parameters[7];  
  beta24 = parameters[8];  
  beta31 = parameters[9];  
  beta32 = parameters[10];  
  beta33 = parameters[11];  
  beta34 = parameters[12];  
  beta41 = parameters[13];  
  beta42 = parameters[14];  
  beta43 = parameters[15];  
  beta44 = parameters[16];  
  gamma1 = parameters[17];  
  gamma2 = parameters[18];  
  gamma3 = parameters[19];  
  gamma4 = parameters[20];  
  
  ## These are the differential equations which describe an S-I-R model  
  ## comprising 4 age groups (in proportion) so all add up to 1.
```

```

## Group 1: Age 0 to 4 years
## Group 2: Age 5 to 19 years
## Group 3: Age 20 to 64 years
## Group 4: Age 65+ years

N1 <- 0.06440
N2 <- 0.20204
N3 <- 0.60074
N4 <- 0.13282

dS1dt = - beta11*S1*(I1/N1) - beta12*S1*(I2/N2) - beta13*S1*(I3/N3) - beta14*S1*(I4/N4)
dS2dt = - beta21*S2*(I1/N1) - beta22*S2*(I2/N2) - beta23*S2*(I3/N3) - beta24*S2*(I4/N4)
dS3dt = - beta31*S3*(I1/N1) - beta32*S3*(I2/N2) - beta33*S3*(I3/N3) - beta34*S3*(I4/N4)
dS4dt = - beta41*S4*(I1/N1) - beta42*S4*(I2/N2) - beta43*S4*(I3/N3) - beta44*S4*(I4/N4)

dI1dt = beta11*S1*(I1/N1) + beta12*S1*(I2/N2) + beta13*S1*(I3/N3) + beta14*S1*(I4/N4) -
gamma1*I1;
dI2dt = beta21*S2*(I1/N1) + beta22*S2*(I2/N2) + beta23*S2*(I3/N3) + beta24*S2*(I4/N4) -
gamma2*I2;
dI3dt = beta31*S3*(I1/N1) + beta32*S3*(I2/N2) + beta33*S3*(I3/N3) + beta34*S3*(I4/N4) -
gamma3*I3;
dI4dt = beta41*S4*(I1/N1) + beta42*S4*(I2/N2) + beta43*S4*(I3/N3) + beta44*S4*(I4/N4) -
gamma4*I4;

dR1dt = gamma1 * I1
dR2dt = gamma2 * I2
dR3dt = gamma3 * I3
dR4dt = gamma4 * I4

return(list(c(dS1dt,dS2dt,dS3dt,dS4dt,dI1dt,dI2dt,dI3dt,dI4dt,dR1dt,dR2dt,dR3dt,dR4dt)));
## This command returns the result, which is the right side of the ODEs as a list,
## back to the solver (i.e. lsoda).
} ## End function specifying the ODEs

#####
## Global variable
##
## Assign numerical values to the parameters and initial conditions of the model
##
## Initial conditions
##
## Demographics proportion of the US Population
## Source: US Census Bureau, ACS Demographic and Housing Estimates,
## 2011 American Community Survey 1-Year Estimates:
##
http://factfinder2.census.gov/faces/tableservices/jsf/pages/productview.xhtml?pid=ACS\_11\_1YR\_DP05&prodType=table
#####

Total_pop <- 310000000 # Approximation of total US population
S1 <- 0.06440 # Proportion of Group 1 in the population (0-4)
S2 <- 0.20204 # Proportion of Group 2 in the population (5-19)
S3 <- 0.60074 # Proportion of Group 3 in the population (20-64)
S4 <- 0.13282 # Proportion of Group 4 in the population (64+)

## Number of infected persons introduced into the USA
seed = 10
## 10 infected person are introduced into the USA (310 million people)

infection_introduction = seed/Total_pop

## Initial proportion of susceptible, infected and recovered populations
## by Group (1, 2, 3, and 4)
S1_entry <- S1
S2_entry <- S2
S3_entry <- S3
S4_entry <- S4
I1_entry <- 0
I2_entry <- 0
I3_entry <- infection_introduction ## Assumption the infected person ("seed")
## is a working adult (Group 3)

```

```

I4_entry <- 0
R1_entry <- 0
R2_entry <- 0
R3_entry <- 0
R4_entry <- 0

## Combine initial conditions into a vector
Y0=c(S1_entry, S2_entry, S3_entry, S4_entry, I1_entry, I2_entry, I3_entry, I4_entry, R1_entry,
R2_entry, R3_entry, R4_entry);

#####
## Recovery rate
## Values for model parameters, units are assumed to be 1/days
## Recovery rate = 0.25; i.e. length of infectiousness = 4 days
gamma_matrix <- c(0.25,0.25,0.25,0.25)
gamma1 <- 0.25;
gamma2 <- 0.25;
gamma3 <- 0.25;
gamma4 <- 0.25;

#####
timevec_interval <- 1 ## "lsoda" output results every day (unit = day)
#####

#####
## Total length of simulation
total_length_simulation <- 365

#####
## The time when the infection was introduced into the USA
## (Number of days since the beginning of the simulation)
time_of_introduction <- 14

#####
## The beginning of the summer break or school closure
## We assume that summer vacation (or school closure) begins 5 days
## after the introduction of the infection
from_intro_to_summer_break <-5
summer_break_start <- time_of_introduction + from_intro_to_summer_break

#####
## Transmission Matrix
#####

## Multiplier to adjust the attack rate and therefore R0
## Epidemiologically, it reflects the overall transmission probability of
## the virus per person-to-person contact
## Assuming 50% of the cases are asymptomatic
## If clinical attack rate ~ 30%, we have "real" AR ~ 60%
## If clinical attack rate ~ 15%, we have "real" AR ~ 30%

for (ARScenario in 1:2){
  if (ARScenario == 1){ #15% Clinical AR
    multiplier <- 0.011047 # CInf == 0.1500055
    CARlabel <- c("CAR15")
  }
  if (ARScenario == 2){ #30% Clinical AR
    multiplier <- 0.016487 # CInf == 0.3000068
    CARlabel <- c("CAR30")
  }
  ## Term time conversational contact matrix obtained from
  ## Eames KTD et al. (2012) Measured Dynamic Social Contact
  ## Patterns Explain the Spread of H1N1v Influenza.
  School_matrix <- matrix(nrow = 4, ncol = 4)
  School_matrix[1,] <- c(4.0196, 1.8137, 7.8039, 0.1373)
  School_matrix[2,] <- c(1.4139, 27.6762, 11.1639, 0.9795)
  School_matrix[3,] <- c(0.8472, 3.8457, 14.7942, 1.0078)
  School_matrix[4,] <- c(0.1048, 0.2857, 6.5673, 2.0980)
  Vacation_matrix <- matrix(nrow = 4, ncol = 4)
  Vacation_matrix[1,] <- c(6.5227, 1.7500, 7.0227, 0.0909)

```

```

Vacation_matrix[2,] <- c(0.9783, 11.5761, 11.7174, 0.5761)
Vacation_matrix[3,] <- c(1.3087, 2.2781, 14.9680, 1.0525)
Vacation_matrix[4,] <- c(0.1442, 0.4279, 5.6512, 1.4326)

n_people <- matrix(nrow = 1, ncol = 4)

B_School_matrix <- matrix(nrow = 4, ncol = 4)
B_Vacation_matrix <- matrix(nrow = 4, ncol = 4)
Contact_matrix <- matrix(nrow = 4, ncol = 4)
Vacation_Contact_matrix <- matrix(nrow = 4, ncol = 4)

experiment <- 3

if (experiment == 1){

  ## Method provided by Dr. John Glasser (part 1) and adapted
  ## from Ken Eames' paper.
  ## Dr. John Glasser's method was applied to convert it to be symmetrical
  Contact_matrix <- sqrt(School_matrix*t(School_matrix))

  # Contact matrix is the square root of A %*% t(A);
  # t(A) is the transpose of A
  # A, i.e. Polymod matrix
  #
  #      [,1]      [,2]      [,3]      [,4]
  # [1,] 4.0196000  1.601371  2.571277  0.1199543
  # [2,] 1.6013714 27.676200  6.552329  0.5290020
  # [3,] 2.5712767  6.552329 14.794200  2.5726494
  # [4,] 0.1199543  0.529002  2.572649  2.0980000

  #> Row_sum # Row sums
  #
  #      [,1]
  # [1,]  8.312202
  # [2,] 36.358902
  # [3,] 26.490455
  # [4,]  5.319606

  #> Mixing_matrix
  #
  #      [,1]      [,2]      [,3]      [,4]
  # [1,] 0.48357821 0.19265308 0.3093376 0.01443111
  # [2,] 0.04404345 0.76119460 0.1802125 0.01454945
  # [3,] 0.09706427 0.24734678 0.5584729 0.09711609
  # [4,] 0.02254948 0.09944384 0.4836166 0.39439013

  #> beta_matrix
  #
  #      [,1]      [,2]      [,3]      [,4]
  # [1,] 0.066271145 0.026401811 0.04239264 0.001977687
  # [2,] 0.026401811 0.456297509 0.10802824 0.008721657
  # [3,] 0.042392640 0.108028242 0.24391198 0.042415271
  # [4,] 0.001977687 0.008721657 0.04241527 0.034589726

  Vacation_Contact_matrix<-sqrt(Vacation_matrix*t(Vacation_matrix))

  # > Vacation_Contact_matrix
  #
  #      [,1]      [,2]      [,3]      [,4]
  # [1,] 6.5227000  1.3084437  3.031601  0.1144892
  # [2,] 1.3084437 11.5761000  5.166566  0.4965009
  # [3,] 3.0316015  5.1665665 14.968000  2.4388292
  # [4,] 0.1144892  0.4965009  2.438829  1.4326000

  # > Vacation_Row_sum
  #
  #      [,1]
  # [1,] 10.977234
  # [2,] 18.547611
  # [3,] 25.604997
  # [4,]  4.482419

  # > Vacation_Mixing_matrix
  #
  #      [,1]      [,2]      [,3]      [,4]
  # [1,] 0.59420249 0.1191961 0.2761717 0.01042970
  # [2,] 0.07054514 0.6241289 0.2785570 0.02676900
  # [3,] 0.11839882 0.2017796 0.5845734 0.09524817

```

```

# [4,] 0.02554183 0.1107663 0.5440877 0.31960419

# > Vacation_beta_matrix
#           [,1]      [,2]      [,3]      [,4]
# [1,] 0.107539755 0.021572312 0.04998201 0.001887584
# [2,] 0.021572312 0.190855161 0.08518118 0.008185811
# [3,] 0.049982013 0.085181181 0.24677742 0.040208977
# [4,] 0.001887584 0.008185811 0.04020898 0.023619276
}

if (experiment == 3){ # Alternative matrix

  n_people[1,] <- c(20067828, 62953784, 187185281, 41385026) # Million. US population.
  for (i in 1:4){
    for (j in 1:4){
      B_School_matrix[i,j] <- n_people[1,i] * School_matrix[i,j] + n_people[1,j] *
School_matrix[j,i]
    }
  }

  for (i in 1:4){
    for (j in 1:4){
      B_Vacation_matrix[i,j] <- n_people[1,i] * Vacation_matrix[i,j] + n_people[1,j] *
Vacation_matrix[j,i]
    }
  }
  for (i in 1:4){
    for (j in 1:4){
      Contact_matrix[i,j] <- B_School_matrix[i,j] / (2*n_people[1,i])
    }
  }
  for (i in 1:4){
    for (j in 1:4){
      Vacation_Contact_matrix[i,j] <- B_Vacation_matrix[i,j] / (2*n_people[1,i])
    }
  }
}

#> Contact_matrix
#           [,1]      [,2]      [,3]      [,4]
# [1,] 4.01960000 3.1245876 7.853134 0.1767123
# [2,] 0.99602730 27.6762000 11.299306 0.5836578
# [3,] 0.84192168 3.8001604 14.794200 1.2298863
# [4,] 0.08568876 0.8878445 5.562800 2.0980000
#> Vacation_Contact_matrix
#           [,1]      [,2]      [,3]      [,4]
# [1,] 6.52270000 2.4094881 9.614885 0.1941388
# [2,] 0.76807445 11.5761000 9.245524 0.4286980
# [3,] 1.03079609 3.1094364 14.968000 1.1509654
# [4,] 0.09413896 0.6521239 5.205839 1.4326000
#> beta_matrix
#           [,1]      [,2]      [,3]      [,4]
# [1,] 0.066271145 0.05151508 0.12947462 0.002913455
# [2,] 0.016421502 0.45629751 0.18629166 0.009622766
# [3,] 0.013880763 0.06265324 0.24391198 0.020277135
# [4,] 0.001412751 0.01463789 0.09171388 0.034589726
#> Vacation_beta_matrix
#           [,1]      [,2]      [,3]      [,4]
# [1,] 0.107539755 0.03972523 0.15852061 0.003200766
# [2,] 0.012663243 0.19085516 0.15243096 0.007067945
# [3,] 0.016994735 0.05126528 0.24677742 0.018975967
# [4,] 0.001552069 0.01075157 0.08582867 0.023619276
}

#####
## School open - matrix ##
#####

Row_sum <- matrix(nrow=4, ncol=1)
for(i in 1:4){

```

```

    Row_sum[i,1]=sum(Contact_matrix[i,1:4])
  }

  Mixing_matrix <- matrix(nrow=4, ncol=4)
  for (i in 1:4){
    Mixing_matrix[i,1:4]<- Contact_matrix[i,1:4]/Row_sum[i,1]
  }

  beta_matrix <- matrix(nrow=4, ncol=4)
  for (i in 1:4){
    for (j in 1:4){
      beta_matrix[i,j] <- Row_sum[i,1] * Mixing_matrix[i,j] * multiplier
    }
  }

  print(beta_matrix)

#####
## Vacation matrix = school closure matrix ##
#####

  Vacation_Row_sum <- matrix(nrow=4, ncol=1)
  for(i in 1:4){
    Vacation_Row_sum[i,1]=sum(Vacation_Contact_matrix[i,1:4])
  }

  Vacation_Mixing_matrix <- matrix(nrow=4, ncol=4)
  for (i in 1:4){
    Vacation_Mixing_matrix[i,1:4]<- Vacation_Contact_matrix[i,1:4]/Vacation_Row_sum[i,1]
  }

  Vacation_beta_matrix <- matrix(nrow=4, ncol=4)
  for (i in 1:4){
    for (j in 1:4){
      Vacation_beta_matrix[i,j] <- Vacation_Row_sum[i,1] * Vacation_Mixing_matrix[i,j] *
multiplier
    }
  }

#####
## Scenarios of School closure ##
#####
for (school_scenario in 1:8){

  ## if baseline, beta_matrix2 == beta_matrix
  if (school_scenario == 1){
    beta_matrix2 <- beta_matrix
    Label <- c("Baseline")
    intervention_length <- 28; # But actually school remains open these 28 d
  }

  ## if we close school, beta_matrix2 == Vacation_beta_matrix
  if (school_scenario > 1){
    beta_matrix2 <- Vacation_beta_matrix
    Label <- c("Closure")
  }

  if (school_scenario == 2) {
    intervention_length <- 7;
    # 7 days; Duration of school closure (1 week)
  }
  if (school_scenario == 3) {
    intervention_length <- 14;
    # 14 days; Duration of school closure (2 weeks)
  }
  if (school_scenario == 4) {
    intervention_length <- 21;
    # 21 days; Duration of school closure (3 weeks)
  }
}

```

```

if (school_scenario == 5) {
  intervention_length <- 28;
  # 28 days; Duration of school closure (4 weeks)
}
if (school_scenario == 6) {
  intervention_length <- 56;
  # 56 days; Duration of school closure (8 weeks)
}
if (school_scenario == 7) {
  intervention_length <- 84;
  # 84 days; Duration of school closure (12 weeks)
}
if (school_scenario == 8) {
  intervention_length <- 140;
  # 140 days; Duration of school closure (20 weeks)
}

intervention_length_reset <- intervention_length
end_of_summer_break = summer_break_start + intervention_length
#reset the time at which intervention ceases

#####
## Create a time series before the introduction of imported cases ##
#####
output <- matrix(0, nrow= (time_of_introduction + 1), ncol=13,dimnames =
list(c(0:time_of_introduction),c("time", "1", "2","3","4","5","6","7","8","9","10","11","12")))
for (n in 1 : (time_of_introduction+1)) {
  output[n, ] <-c((n-1), S1, S2, S3, S4, 0,0,0,0,0,0,0,0)
}

ScenLabel <- paste("P_",Label,sep="")

#####
## Introduction of imported cases. Before summer break ##
#####
timevec=seq(time_of_introduction,summer_break_start,by=1);
#this creates a vector of times for which integration is evaluated

parvec=c(beta_matrix[1,1], beta_matrix[1,2], beta_matrix[1,3], beta_matrix[1,4],
beta_matrix[2,1], beta_matrix[2,2], beta_matrix[2,3], beta_matrix[2,4], beta_matrix[3,1],
beta_matrix[3,2], beta_matrix[3,3], beta_matrix[3,4], beta_matrix[4,1], beta_matrix[4,2],
beta_matrix[4,3], beta_matrix[4,4], gamma1, gamma2, gamma3, gamma4);
# This combines all parameters into a vector called parvec
# which is sent to the ODE function

odeoutput=ode(y=Y0, times=timevec, func=odeequations, parms=parvec);
pre_intervention = odeoutput[length(odeoutput[,1]),1]

# Variable value at the beginning of summer break (or school closure)
S1_break <- odeoutput[length(odeoutput[,1]),2];
S2_break <- odeoutput[length(odeoutput[,1]),3];
S3_break <- odeoutput[length(odeoutput[,1]),4];
S4_break <- odeoutput[length(odeoutput[,1]),5];
I1_break <- odeoutput[length(odeoutput[,1]),6];
I2_break <- odeoutput[length(odeoutput[,1]),7];
I3_break <- odeoutput[length(odeoutput[,1]),8];
I4_break <- odeoutput[length(odeoutput[,1]),9];
R1_break <- odeoutput[length(odeoutput[,1]),10];
R2_break <- odeoutput[length(odeoutput[,1]),11];
R3_break <- odeoutput[length(odeoutput[,1]),12];
R4_break <- odeoutput[length(odeoutput[,1]),13];

## CInf stands for cumulative number of infections,
## i.e. cumulative attack rates, at the beginning of summer break
CInf1_break <- S1_entry - S1_break
CInf2_break <- S2_entry - S2_break
CInf3_break <- S3_entry - S3_break
CInf4_break <- S4_entry - S4_break

```



```
#####
## Summer break (School closure)

Y1=c(S1_break, S2_break, S3_break, S4_break, I1_break, I2_break, I3_break, I4_break, R1_break,
R2_break, R3_break, R4_break);
intervention_length = intervention_length_reset

timevec1=seq( pre_intervention, end_of_summer_break, by=timevec_interval);
## This creates a vector of times for which integration is evaluated

parvec1=c(beta_matrix2[1,1], beta_matrix2[1,2], beta_matrix2[1,3], beta_matrix2[1,4],
beta_matrix2[2,1], beta_matrix2[2,2], beta_matrix2[2,3], beta_matrix2[2,4], beta_matrix2[3,1],
beta_matrix2[3,2], beta_matrix2[3,3], beta_matrix2[3,4], beta_matrix2[4,1], beta_matrix2[4,2],
beta_matrix2[4,3], beta_matrix2[4,4], gamma1, gamma2, gamma3, gamma4);
## This combines all parameters into a vector called parvec which is sent to the ODE function

odeoutput1=ode(y=Y1, times=timevec1, func=odeequations, parms=parvec1);

intervention_end = odeoutput1[length(odeoutput1[,1]),1];

S1_break_end <- odeoutput1[length(odeoutput1[,1]),2];
S2_break_end <- odeoutput1[length(odeoutput1[,1]),3];
S3_break_end <- odeoutput1[length(odeoutput1[,1]),4];
S4_break_end <- odeoutput1[length(odeoutput1[,1]),5];
I1_break_end <- odeoutput1[length(odeoutput1[,1]),6];
I2_break_end <- odeoutput1[length(odeoutput1[,1]),7];
I3_break_end <- odeoutput1[length(odeoutput1[,1]),8];
I4_break_end <- odeoutput1[length(odeoutput1[,1]),9];
R1_break_end <- odeoutput1[length(odeoutput1[,1]),10];
R2_break_end <- odeoutput1[length(odeoutput1[,1]),11];
R3_break_end <- odeoutput1[length(odeoutput1[,1]),12];
R4_break_end <- odeoutput1[length(odeoutput1[,1]),13];

## CInf stands for cumulative number of infections, i.e. cumulative attack rates, at the
beginning of summer break
CInf1_break_end <- S1_entry - S1_break_end
CInf2_break_end <- S2_entry - S2_break_end
CInf3_break_end <- S3_entry - S3_break_end
CInf4_break_end <- S4_entry - S4_break_end

#####
## School starts in Fall (when schools re-open) ##
#####

Y2=c(S1_break_end, S2_break_end, S3_break_end, S4_break_end, I1_break_end, I2_break_end,
I3_break_end, I4_break_end, R1_break_end, R2_break_end, R3_break_end, R4_break_end);

timevec2=seq(end_of_summer_break, total_length_simulation, by=timevec_interval);
# This creates a vector of times for which integration is evaluated

parvec2=c(beta_matrix[1,1], beta_matrix[1,2], beta_matrix[1,3], beta_matrix[1,4],
beta_matrix[2,1], beta_matrix[2,2], beta_matrix[2,3], beta_matrix[2,4], beta_matrix[3,1],
beta_matrix[3,2], beta_matrix[3,3], beta_matrix[3,4], beta_matrix[4,1], beta_matrix[4,2],
beta_matrix[4,3], beta_matrix[4,4], gamma1, gamma2, gamma3, gamma4);
# This combines all parameters into a vector called parvec which is sent to the ODE function

odeoutput2=ode(y=Y2, times=timevec2, func=odeequations, parms=parvec2);

intervention_end = odeoutput2[length(odeoutput2[,1]),1];

S1_end <- odeoutput2[length(odeoutput2[,1]),2];
S2_end <- odeoutput2[length(odeoutput2[,1]),3];
S3_end <- odeoutput2[length(odeoutput2[,1]),4];
S4_end <- odeoutput2[length(odeoutput2[,1]),5];
I1_end <- odeoutput2[length(odeoutput2[,1]),6];
I2_end <- odeoutput2[length(odeoutput2[,1]),7];
I3_end <- odeoutput2[length(odeoutput2[,1]),8];
I4_end <- odeoutput2[length(odeoutput2[,1]),9];
R1_end <- odeoutput2[length(odeoutput2[,1]),10];
```

```

R2_end <- odeoutput2[length(odeoutput2[,1]),11];
R3_end <- odeoutput2[length(odeoutput2[,1]),12];
R4_end <- odeoutput2[length(odeoutput2[,1]),13];

outputall <- matrix(0,nrow=366,ncol=13)
outputall[1:(time_of_introduction+1),] <- output[1:length(output[,1]),]
outputall[(time_of_introduction+1):(summer_break_start+1),] <-
odeoutput[1:length(odeoutput[,1]),]
outputall[(summer_break_start+1):(end_of_summer_break+1),] <-
odeoutput1[1:length(odeoutput1[,1]),]
outputall[(end_of_summer_break+1):366,] <- odeoutput2[1:length(odeoutput2[,1]),]
## Convert to numbers
newoutputall <- outputall
newoutputall[,2:13] <- Total_pop*outputall[,2:13]

#####
## Assumption of symptomatic to asymptomatic ratio
## If we want to count all cases (symptomatic and asymptomatic), or
## if we assume that all cases are symptomatic, then sym_ratio = 1
## If we assume that only 1 in 2 cases are symptomatic, then sym_ratio = 0.5
## If we assume that only 1 in 3 cases are symptomatic, then sym_ratio = 0.33
sym_ratio <- 0.5

countall <- matrix(0,nrow=366,ncol=7)
countall[,1] <- newoutputall[,1]
countall[,2:5] <- newoutputall[,6:9]*sym_ratio
# Divided by sym_ratio
for (n in 1:366) {
  countall[n,6] <- (sum(newoutputall[n,6:9]))*sym_ratio
  # Total incidence on a given day
  # Divided by sym_ratio
  countall[n,7] <- (Total_pop - sum(newoutputall[n,2:5]))*sym_ratio
  # Count cumulative incidence
  # i.e. Total population - Total Susceptible population
  # Divided by sym_ratio
}

filename_countall <- paste(experiment, CARlabel, ScenLabel, seed, intervention_length,
"incidence.csv", sep="_")
write.csv(countall, filename_countall)

## CInf stands for cumulative number of infections, i.e. cumulative attack rates
## Divided by 2 (sym_ratio = 0.5): Assumption: 1 in 2 infected persons is asymptomatic

CInf1 <- (S1_entry - S1_end)* sym_ratio
CInf2 <- (S2_entry - S2_end)* sym_ratio
CInf3 <- (S3_entry - S3_end)* sym_ratio
CInf4 <- (S4_entry - S4_end)* sym_ratio
CInf <- CInf1 + CInf2 + CInf3 + CInf4

## Write data file
Data <- matrix(0, nrow=3, ncol=1)
peak <- which.max(countall[,6])
print(sprintf("Peak: Day %s",countall[peak,1]))
Data[1,1] <- countall[peak,1]
print(sprintf("Daily number of new cases at peak time: %s",countall[peak,6]))
Data[2,1] <- countall[peak,6]
Data[3,1] <- CInf
print(CInf)
filename_csv <- paste(experiment,CARlabel,ScenLabel,seed,intervention_length,"data.csv",sep="_")
write.csv(Data, filename_csv)
}
}

```

## R code 2: “Flu\_School\_Closure\_CID\_Data.r”

## This R code creates the data summary files. This is optional for our modeling purpose.

```
experiment <- 3 ## Type 1 for Main analysis; 3 for Alternative matrix

filename_M1 <- paste(experiment, "_CAR30_P_Baseline_10_28_data.csv", sep = "")
filename_M2 <- paste(experiment, "_CAR30_P_Closure_10_7_data.csv", sep = "")
filename_M3 <- paste(experiment, "_CAR30_P_Closure_10_14_data.csv", sep = "")
filename_M4 <- paste(experiment, "_CAR30_P_Closure_10_21_data.csv", sep = "")
filename_M5 <- paste(experiment, "_CAR30_P_Closure_10_28_data.csv", sep = "")
filename_M6 <- paste(experiment, "_CAR30_P_Closure_10_56_data.csv", sep = "")
filename_M7 <- paste(experiment, "_CAR30_P_Closure_10_84_data.csv", sep = "")
filename_M8 <- paste(experiment, "_CAR30_P_Closure_10_140_data.csv", sep = "")

filename_M9 <- paste(experiment, "_CAR15_P_Baseline_10_28_data.csv", sep = "")
filename_M10 <- paste(experiment, "_CAR15_P_Closure_10_7_data.csv", sep = "")
filename_M11 <- paste(experiment, "_CAR15_P_Closure_10_14_data.csv", sep = "")
filename_M12 <- paste(experiment, "_CAR15_P_Closure_10_21_data.csv", sep = "")
filename_M13 <- paste(experiment, "_CAR15_P_Closure_10_28_data.csv", sep = "")
filename_M14 <- paste(experiment, "_CAR15_P_Closure_10_56_data.csv", sep = "")
filename_M15 <- paste(experiment, "_CAR15_P_Closure_10_84_data.csv", sep = "")
filename_M16 <- paste(experiment, "_CAR15_P_Closure_10_140_data.csv", sep = "")

M1 <- read.csv(filename_M1)
M2 <- read.csv(filename_M2)
M3 <- read.csv(filename_M3)
M4 <- read.csv(filename_M4)
M5 <- read.csv(filename_M5)
M6 <- read.csv(filename_M6)
M7 <- read.csv(filename_M7)
M8 <- read.csv(filename_M8)

M9 <- read.csv(filename_M9)
M10 <- read.csv(filename_M10)
M11 <- read.csv(filename_M11)
M12 <- read.csv(filename_M12)
M13 <- read.csv(filename_M13)
M14 <- read.csv(filename_M14)
M15 <- read.csv(filename_M15)
M16 <- read.csv(filename_M16)

DataMatrix <- matrix(nrow=length(M8[,1]), ncol=16)
DataMatrix[,1] <- M1[,2]
DataMatrix[,2] <- M2[,2]
DataMatrix[,3] <- M3[,2]
DataMatrix[,4] <- M4[,2]
DataMatrix[,5] <- M5[,2]
DataMatrix[,6] <- M6[,2]
DataMatrix[,7] <- M7[,2]
DataMatrix[,8] <- M8[,2]
DataMatrix[,9] <- M9[,2]
DataMatrix[,10] <- M10[,2]
DataMatrix[,11] <- M11[,2]
DataMatrix[,12] <- M12[,2]
DataMatrix[,13] <- M13[,2]
DataMatrix[,14] <- M14[,2]
DataMatrix[,15] <- M15[,2]
DataMatrix[,16] <- M16[,2]

Report <- matrix(nrow=3, ncol=16)
Report[1,] <- DataMatrix[1,]
Report[2,] <- DataMatrix[2,]
Report[3,] <- DataMatrix[3,]

# Write datafile
filename_csv <- paste(experiment, "CAR30_report_data.csv", sep = "_")
write.csv(Report[,1:8], filename_csv)
filename_csv2 <- paste(experiment, "CAR15_report_data.csv", sep = "_")
write.csv(Report[,9:16], filename_csv2)
```

## R code 3: “Flu\_School\_Closure\_CID\_Plot.r”

```
## This R code was used to create the figures in the article.

AR <- c("_CAR15")
## Type "_CAR15" for cumulative attack rate 15% scenario
## Type "_CAR30" for cumulative attack rate 30% scenario

## Read the data files
filename_M1 <- paste(experiment,AR,"_P_Baseline_10_28_incidence.csv", sep = "")
filename_M2 <- paste(experiment,AR,"_P_Closure_10_7_incidence.csv", sep = "")
filename_M3 <- paste(experiment,AR,"_P_Closure_10_14_incidence.csv", sep = "")
filename_M4 <- paste(experiment,AR,"_P_Closure_10_21_incidence.csv", sep = "")
filename_M5 <- paste(experiment,AR,"_P_Closure_10_28_incidence.csv", sep = "")
filename_M6 <- paste(experiment,AR,"_P_Closure_10_56_incidence.csv", sep = "")
filename_M7 <- paste(experiment,AR,"_P_Closure_10_84_incidence.csv", sep = "")
filename_M8 <- paste(experiment,AR,"_P_Closure_10_140_incidence.csv", sep = "")
M1 <- read.csv(filename_M1)
M2 <- read.csv(filename_M2)
M3 <- read.csv(filename_M3)
M4 <- read.csv(filename_M4)
M5 <- read.csv(filename_M5)
M6 <- read.csv(filename_M6)
M7 <- read.csv(filename_M7)
M8 <- read.csv(filename_M8)

## Plot the figures
plot(M1[,2],M1[,7],type="l",xlab="Time (days)",ylab="Cases
(Million)",col="black",lwd=3,lty=1,xlim=c(0,365),ylim=c(0,25000000), yaxt='n')
  ## Suppress the default y-axis with yaxt='n'
  lines(M2[,2],M2[,7],type="l",col="green",lwd=3,lty=2)
  lines(M3[,2],M3[,7],type="l",col="light blue",lwd=3,lty=3)
  lines(M4[,2],M4[,7],type="l",col="gold",lwd=3,lty=1)
  lines(M5[,2],M5[,7],type="l",col="red",lwd=3,lty=2)
  lines(M6[,2],M6[,7],type="l",col="blue",lwd=3,lty=3)
  lines(M7[,2],M7[,7],type="l",col="magenta",lwd=3,lty=4)
  lines(M8[,2],M8[,7],type="l",col="grey",lwd=3,lty=5)

## Add the tailor-made y-axis
if (experiment == 1){ axis(2, at = c(0,4000000,10000000,16000000), labels=c("0","4","10","16"))}
if (experiment == 3){ axis(2, at = c(0,3000000,10000000,14000000), labels=c("0","3","10","14"))}

## Add arrows
arrows(summer_break_start,11000000,(summer_break_start+7),11000000,code=3,col=c("green"))
arrows(summer_break_start,10000000,(summer_break_start+14),10000000,code=3,col=c("light
blue"))
arrows(summer_break_start,9000000,(summer_break_start+21),9000000,code=3,col=c("gold"))
arrows(summer_break_start,8000000,(summer_break_start+28),8000000,code=3,col=c("red"))
arrows(summer_break_start,7000000,(summer_break_start+56),7000000,code=3,col=c("blue"))
arrows(summer_break_start,6000000,(summer_break_start+84),6000000,code=3,col=c("magenta"))
arrows(summer_break_start,5000000,(summer_break_start+140),5000000,code=3,col=c("grey"))
arrows(time_of_introduction,5000000,time_of_introduction,0,code=2,col=c("purple"))

## Add legend
legend(0,25000000,c("Baseline","Closure 7d", "Closure 14d","Closure 21d","Closure
28d","Closure 56d","Closure 84d","Closure 140d"), col = c("black","green","light
blue","gold","red","blue","magenta","grey"), lty=c(1,2,3,1,2,3,4,5), lwd = 3, ncol=3)
```

## R code 4: “Flu\_School\_Closure\_CID\_Difference\_eqn.r”

```
#####
## Difference Equation SIR Model for School Closure in an Influenza Pandemic ##
#####
## Corresponding to the Ordinary Differential Equation SIR Model          ##
## for School Closure in an Influenza Pandemic                          ##
#####
## Written by Isaac Chun-Hai FUNG, PhD                                   ##
## June 11, 2013; revised on Sep 12, 2013;                               ##
## edited for publication on Nov 21, 2014                                ##
#####

#####
## Time step ##
#####
for (experiment in 1:4){

  # Time step size for difference equation model
  if (experiment == 1) {Timestep_size <- 1}
  if (experiment == 2) {Timestep_size <- 0.1}
  if (experiment == 3) {Timestep_size <- 0.01}
  if (experiment == 4) {Timestep_size <- 0.001}

  # Duration (days) of simulation
  Simulation_length <- 365

  Total_timesteps <- Simulation_length / Timestep_size

  #####
  ## School closure time frame ##
  #####

  school_close <- 5 / Timestep_size
  school_closure_length <- 28 / Timestep_size
  school_reopen <- school_close + school_closure_length
  print(school_close)
  print(school_reopen)

  #####
  ## Assumption of symptomatic to asymptomatic ratio                    ##
  ## If we want to count all cases (symptomatic and asymptomatic), or   ##
  ## if we assume that all cases are symptomatic, then sym_ratio = 1    ##
  ## If we assume that only 1 in 2 cases are symptomatic, then sym_ratio = 0.5 ##
  ## If we assume that only 1 in 3 cases are symptomatic, then sym_ratio = 0.33 ##
  #####
  sym_ratio <- 0.5

  #####
  ## Transmission matrices ##
  #####
  multiplier <- 0.016487 # CInf == 0.2999887
  CARlabel <- c("CAR30")

  ## Rate of Recovery
  gamma_matrix <- c(0.25,0.25,0.25,0.25)

  #####
  ## School open - matrix ##
  #####
  ## School matrix (term time) obtained from Eames KTD et al. (2012) Measured
  ## Dynamic Social Contact Patterns Explain the Spread of H1N1v Influenza.
  School_matrix <- matrix(nrow = 4, ncol = 4)
  School_matrix[1,] <- c(4.0196, 1.8137, 7.8039, 0.1373)
  School_matrix[2,] <- c(1.4139, 27.6762, 11.1639, 0.9795)
  School_matrix[3,] <- c(0.8472, 3.8457, 14.7942, 1.0078)
  School_matrix[4,] <- c(0.1048, 0.2857, 6.5673, 2.0980)

  Contact_matrix <- matrix(nrow = 4, ncol = 4)
  Contact_matrix <- sqrt(School_matrix*t(School_matrix))
}
```

```

# Contact matrix is the square root of A %*% t(A); t(A) is the transpose of A
# A, i.e. Polymod matrix
#           [,1]      [,2]      [,3]      [,4]
# [1,] 4.0196000  1.601371  2.571277  0.1199543
# [2,] 1.6013714 27.676200  6.552329  0.5290020
# [3,] 2.5712767  6.552329 14.794200  2.5726494
# [4,] 0.1199543  0.529002  2.572649  2.0980000

Row_sum <- matrix(nrow=4, ncol=1)
for(i in 1:4){
  Row_sum[i,1]=sum(Contact_matrix[i,1:4])
}
#> Row_sum # Row sums
#           [,1]
# [1,]  8.312202
# [2,] 36.358902
# [3,] 26.490455
# [4,]  5.319606

Mixing_matrix <- matrix(nrow=4, ncol=4)
for (i in 1:4){
  Mixing_matrix[i,1:4]<- Contact_matrix[i,1:4]/Row_sum[i,1]
}
#>   Mixing_matrix
#           [,1]      [,2]      [,3]      [,4]
# [1,] 0.48357821 0.19265308 0.3093376 0.01443111
# [2,] 0.04404345 0.76119460 0.1802125 0.01454945
# [3,] 0.09706427 0.24734678 0.5584729 0.09711609
# [4,] 0.02254948 0.09944384 0.4836166 0.39439013
beta_matrix <- matrix(nrow=4, ncol=4)
for (i in 1:4){
  for (j in 1:4){
    beta_matrix[i,j] <- Row_sum[i,1] * Mixing_matrix[i,j] * multiplier
  }
}
R0_matrix <- matrix(nrow=1, ncol=4)
for (j in 1:4) {
  R0_matrix[,j] <- sum(beta_matrix[,j]) / gamma_matrix[j]
}

#####
## Vacation matrix = school closure matrix ##
#####
Vacation_matrix <- matrix(nrow = 4, ncol = 4)
Vacation_matrix[1,] <- c(6.5227, 1.7500, 7.0227, 0.0909)
Vacation_matrix[2,] <- c(0.9783, 11.5761, 11.7174, 0.5761)
Vacation_matrix[3,] <- c(1.3087, 2.2781, 14.9680, 1.0525)
Vacation_matrix[4,] <- c(0.1442, 0.4279, 5.6512, 1.4326)

Vacation_Contact_matrix <- matrix(nrow = 4, ncol = 4)
Vacation_Contact_matrix <- sqrt(Vacation_matrix*t(Vacation_matrix))
# > Vacation_Contact_matrix
#           [,1]      [,2]      [,3]      [,4]
# [1,] 6.5227000  1.3084437  3.031601 0.1144892
# [2,] 1.3084437 11.5761000  5.166566 0.4965009
# [3,] 3.0316015  5.1665665 14.968000 2.4388292
# [4,] 0.1144892  0.4965009  2.438829 1.4326000

Vacation_Row_sum <- matrix(nrow=4, ncol=1)
for(i in 1:4){
  Vacation_Row_sum[i,1]=sum(Vacation_Contact_matrix[i,1:4])
}
# > Vacation_Row_sum
#           [,1]
# [1,] 10.977234
# [2,] 18.547611
# [3,] 25.604997
# [4,]  4.482419

Vacation_Mixing_matrix <- matrix(nrow=4, ncol=4)
for (i in 1:4){

```

```

    Vacation_Mixing_matrix[i,1:4] <- Vacation_Contact_matrix[i,1:4]/Vacation_Row_sum[i,1]
  }
  # > Vacation_Mixing_matrix
  #      [,1]      [,2]      [,3]      [,4]
# [1,] 0.59420249 0.1191961 0.2761717 0.01042970
# [2,] 0.07054514 0.6241289 0.2785570 0.02676900
# [3,] 0.11839882 0.2017796 0.5845734 0.09524817
# [4,] 0.02554183 0.1107663 0.5440877 0.31960419

Vacation_beta_matrix <- matrix(nrow=4, ncol=4)
for (i in 1:4){
  for (j in 1:4){
    Vacation_beta_matrix[i,j] <- Vacation_Row_sum[i,1] * Vacation_Mixing_matrix[i,j] *
multiplier
  }
}
Vacation_R0_matrix <- matrix(nrow = 1, ncol = 4)
for (j in 1:4) {
  Vacation_R0_matrix[,j] <- sum(Vacation_beta_matrix[,j]) / gamma_matrix[j]
}

#####
## Equations ##
#####

## These are the difference equations which describe an S-I-R model
## comprising children and adults as separated groups
## Group 1: Age 0 - 4 years
## Group 2: Age 5 - 19 years
## Group 3: Age 20 - 64 years
## Group 4: Age 65+ years

Total_pop <- 310000000 ## Approximation of total US population
Pop1 <- 0.06440 * Total_pop ## Proportion of Group 1 in the population (0-4)
Pop2 <- 0.20204 * Total_pop ## Proportion of Group 2 in the population (5-19)
Pop3 <- 0.60074 * Total_pop ## Proportion of Group 3 in the population (20-64)
Pop4 <- 0.13282 * Total_pop ## Proportion of Group 4 in the population (64+)

## Initial number of infected people coming from overseas
seed = 10

## Create a matrix for the variables
y <- matrix(nrow = Total_timesteps+1, ncol = 13)

## Initial proportion of susceptible, infected and recovered populations
## by Group (1, 2, 3, and 4)
y_initial <- matrix(0,1,13)
y_initial[1,1:13] <- c(0,Pop1,Pop2,Pop3,Pop4,0,0,seed,0,0,0,0,0)
## Assumption: the infected person ("seed") is a working adult (Group 3)

y[1,] <- y_initial[1,]
S1 <- y[1,2]
S2 <- y[1,3]
S3 <- y[1,4]
S4 <- y[1,5]
I1 <- y[1,6]
I2 <- y[1,7]
I3 <- y[1,8]
I4 <- y[1,9]
R1 <- y[1,10]
R2 <- y[1,11]
R3 <- y[1,12]
R4 <- y[1,13]

N1 <- 0.06440 * Total_pop
N2 <- 0.20204 * Total_pop
N3 <- 0.60074 * Total_pop
N4 <- 0.13282 * Total_pop

gamma1 = gamma_matrix[1]
gamma2 = gamma_matrix[2]

```

```

gamma3 = gamma_matrix[3]
gamma4 = gamma_matrix[4]

b <- beta_matrix

for (timestep in 1:Total_timesteps){

  ## Apply the appropriate beta matrix at the right time
  if (timestep >= school_close && timestep < school_reopen) { b <- Vacation_beta_matrix }
  else { b <- beta_matrix }

  ## These are the equations for the S-I-R model
  ## Susceptible populations
  S1_update = S1 - Timestep_size * (b[1,1]*S1*(I1/N1) + b[1,2]*S1*(I2/N2) +
b[1,3]*S1*(I3/N3) + b[1,4]*S1*(I4/N4))
  S2_update = S2 - Timestep_size * (b[2,1]*S2*(I1/N1) + b[2,2]*S2*(I2/N2) +
b[2,3]*S2*(I3/N3) + b[2,4]*S2*(I4/N4))
  S3_update = S3 - Timestep_size * (b[3,1]*S3*(I1/N1) + b[3,2]*S3*(I2/N2) +
b[3,3]*S3*(I3/N3) + b[3,4]*S3*(I4/N4))
  S4_update = S4 - Timestep_size * (b[4,1]*S4*(I1/N1) + b[4,2]*S4*(I2/N2) +
b[4,3]*S4*(I3/N3) + b[4,4]*S4*(I4/N4))

  ## Infected(Infectious) populations
  I1_update = I1 + Timestep_size * (b[1,1]*S1*(I1/N1) + b[1,2]*S1*(I2/N2) +
b[1,3]*S1*(I3/N3) + b[1,4]*S1*(I4/N4) - gamma1*I1)
  I2_update = I2 + Timestep_size * (b[2,1]*S2*(I1/N1) + b[2,2]*S2*(I2/N2) +
b[2,3]*S2*(I3/N3) + b[2,4]*S2*(I4/N4) - gamma2*I2)
  I3_update = I3 + Timestep_size * (b[3,1]*S3*(I1/N1) + b[3,2]*S3*(I2/N2) +
b[3,3]*S3*(I3/N3) + b[3,4]*S3*(I4/N4) - gamma3*I3)
  I4_update = I4 + Timestep_size * (b[4,1]*S4*(I1/N1) + b[4,2]*S4*(I2/N2) +
b[4,3]*S4*(I3/N3) + b[4,4]*S4*(I4/N4) - gamma4*I4)

  ## Recovered populations
  R1_update = R1 + Timestep_size * gamma1 * I1
  R2_update = R2 + Timestep_size * gamma2 * I2
  R3_update = R3 + Timestep_size * gamma3 * I3
  R4_update = R4 + Timestep_size * gamma4 * I4

  ## Update the matrix
  y[timestep+1,1]<- y[timestep,1] + Timestep_size

  y[timestep+1,2]<- S1_update;
  y[timestep+1,3]<- S2_update;
  y[timestep+1,4]<- S3_update;
  y[timestep+1,5]<- S4_update;      # Susceptible

  y[timestep+1,6]<- I1_update;
  y[timestep+1,7]<- I2_update;
  y[timestep+1,8]<- I3_update;
  y[timestep+1,9]<- I4_update;      # Infected (Infectious)

  y[timestep+1,10]<- R1_update;
  y[timestep+1,11]<- R2_update;
  y[timestep+1,12]<- R3_update;
  y[timestep+1,13]<- R4_update;      # Recovered

  ## Update the state variable for the next time step
  S1 <- S1_update;
  S2 <- S2_update;
  S3 <- S3_update;
  S4 <- S4_update;      # Susceptible

  I1 <- I1_update;
  I2 <- I2_update;
  I3 <- I3_update;
  I4 <- I4_update;      # Infected (Infectious)

  R1 <- R1_update;
  R2 <- R2_update;
  R3 <- R3_update;
  R4 <- R4_update;      # Recovered

```



```

    }
newoutputall <- y

newmatrix <- matrix(ncol = 2, nrow = length(newoutputall[,1]) )
newmatrix[,1] <- y[,1]
for (n in 1:length(newoutputall[,1])){
  newmatrix[n,2] <- sum(newoutputall[n,6:9])*sym_ratio
}

## Plot figures
plot(newmatrix[,1], newmatrix[,2], type="l", xlab="Time (days)", ylab="Cases (million)",
col="red", lwd=3, lty=1, xlim=c(0,100), ylim=c(0,15000000), yaxt='n')

## Add y-axis, arrows and text
axis(2, at = c(0,13000000), labels=c("0","13"))
arrows(0,5000000,0,0,code=2,col=c("black"))
arrows(5,8000000,5,0,code=2,col=c("dark grey"))
arrows(5,8000000,(5+28),8000000,code=2,col=c("dark grey"))
text(10,6000000, "outbreak begins")
text(18,9000000,"school closure (28d)")
print(max(newmatrix[,2]))
}

```