Carbapenemase-producing Klebsiella pneumoniae (CPKP) diffusion on a network

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Abstract

The studies on the mathematical diffusion model of disease are very important to plan a strategy to avoid the contagion. These kind of models start by the SIS model and can be improved, an example is given by the model for the diffusion of the Carbapenemase-producing Klebsiella pneumoniae (CPKP) in a surgical unit. For this purpose the network study can be fundamental to understand the particular behaviour of the diffusion in a discreet system instead of the mean field approximation. This brief work try to seek this purpose.

Contents

Chapter 1

SIS model for epidemic diffusion

Diffusion of pathologies can be modelled using the SIS model which is a model involving two states. It can be easily implemented on a network by the fact that the network could be seen as a numerical simulation of the spread in a real network.

1.1 SIS model features

The SIS (Susceptible-Infected-Susceptible) is based on the idea that all the elements which are attacked by the pathology can have two possible state: susceptible and infected. If a susceptible elements is touched by a infected one than with a possibility λ the first one become infected:

$$S(i) + I(j) \xrightarrow{\lambda} I(i) + I(j). \tag{1.1}$$

There could be also a probability for the inverse process μ . We can make a mean field description by solving this system of differential equations:

$$\frac{ds(t)}{dt} = \mu \rho(t) - \lambda \langle k \rangle \rho(t) s(t)
\frac{d\rho(t)}{dt} = -\frac{ds(t)}{dt}$$
(1.2)

in the condition of an isolated system and considering the initial fraction of infected nodes very little. $\langle k \rangle$ is the average number of contacts.

Using this equation we can get a estimation of the epidemic threshold by considering a steady state for ρ :

$$\mu \rho(t) - \lambda \langle k \rangle \rho(t) (1 - \rho(t)) = 0$$

$$\rho = 1 - \frac{\mu}{\lambda \langle k \rangle}$$
(1.3)

so we can define two cases:

$$\lambda < \frac{\mu}{\langle k \rangle} \Rightarrow \rho = 0 \quad No \ epidemic$$

$$\lambda > \frac{\mu}{\langle k \rangle} \Rightarrow \rho = 1 - \frac{\mu}{\lambda \langle k \rangle} \quad Endemic \ state$$
(1.4)

This can be seen in the figure 1.1 where one can observe the phase transition with order parameter ρ .

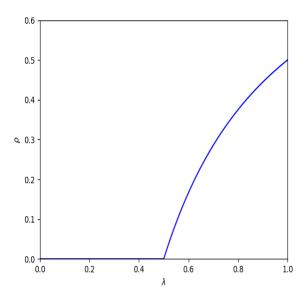


Figure 1.1: Typical behaviour of an order parameter in presence of a phase transition.

If we want to introduce a way to decrease the contagion we can add an immunization effect which decrease the transmission probability λ by a factor (1-g). By doing this we can define a critical value for g over which we stop the contagion g_c .

1.2 SIS model for heterogeneous sample

 $\langle k \rangle$ is a mean quantity but on the network the connectivity could follow a distribution p(k)dk, following this idea we can model the SIS diffusion on a heterogeneous sample. For an heterogeneous sample we have a differential equation for each degree k:

$$\frac{ds_k(t)}{dt} = \mu \rho_k(t) - \lambda k s_k(t) \Theta(t)$$

$$\frac{d\rho_k(t)}{dt} = -\frac{ds_k(t)}{dt}$$
(1.5)

where:

$$\Theta = \frac{\sum_{k} kp(k)\rho_{k}(t)}{\sum_{k} kp(k)}$$
(1.6)

is the weighted average parameter

There is a big difference between the mean system and the heterogeneous system that can be seen by analyzing Θ , in fact we obtain:

$$\Theta = \frac{\sum_{k} kp(k) \frac{\lambda k\Theta}{1 + \lambda k\Theta}}{\sum_{k} kp(k)}$$
 (1.7)

so $\Theta = 0$ implies $\rho = 0$, the solution $\Theta > 0$ is given by:

$$\frac{d}{d\Theta} \left(\frac{\sum_{k} kp(k) \frac{\lambda k\Theta}{1 + \lambda k\Theta}}{\sum_{k} kp(k)} \right) |_{\Theta=0} \ge 1$$
(1.8)

so the critical value for λ will be: $\langle k \rangle / \langle k^2 \rangle$. This means that if the second momentum of the distribution diverges than the epidemic threshold goes to zero replacing the discontinuity of the figure 1.1 with a smooth function.

1.3 SIS model on network

The SIS model can be implemented on a network using a Markov chain. The following code is based on an event driven algorithm.

```
import math
2 import random
3 from networkx import *
4 from numpy import *
6 #Preparing the graph
7 G=fast_gnp_random_graph(20, 0.5, seed=None, directed=False)
8 State=['Susceptible']
9 set_node_attributes(G, State, 'State')
#Preparing the infection source
rand=random.randint(0,19)
G.nodes[0]['State']=['Infected']
14 infection_list=[]
infection_list.append(rand)
17 #Parameters
18 lambd=0.1
19 \text{ mu} = 0.1
_{20} Time = 10
21
22 #Infection
  for t in range(Time):
24
    #Decontamination
      for node in G.nodes():
26
           if G.nodes[node]['State'] == ['Infected']:
27
               rand_head=random.uniform(0,1)
28
               if rand_head <= mu:</pre>
29
                   G.nodes[node]['State']=['Susceptible']
30
31
      for node in G.nodes():
           if G.nodes[node]['State'] == ['Infected']:
               neighbours=list(G.neighbors(i))
34
               for j in range(len(neighbours)):
                   if Node_state[neighbours[j]] == ['Susceptible']:
                        rand_inf=random.uniform(0,1)
37
                        if rand_inf <= lambd:</pre>
38
                            infection_list.append(neighbours[j])
39
      infect_list=list(set(infect_list)) #Keeping only the unique value
      for each infected node
      for i in range(len(infection_list)):
41
           G.nodes[infection_list[i]]['State']=['Infected']
42
```

Chapter 2

CPKP diffusion

In 2010 Vana Sypsa, Mina Psichogiou, Georgia-Aikaterina Bouzala, Linos Hadjihannas, Angelos Hatzakis1, Georgios L. Daikos conduced a study abouth the diffusion of the carbapenemase-producing Klebsiella pneumoniae (CPKP) in a surgical unit in order to understand the behaviour and the possible way to control the infection.

2.1 The model features

The model include two possible role: the HCW (health care worker) and the patient. The model is based on the scheme in figure 2.1.

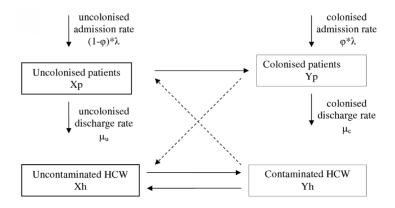


Figure 2.1: Scheme of CPKP diffusion from the article.

The control model scheme is shown by the figure 2.2.

I have decided to implement the model including the possibility of a direct contagion between the HCW to get a more general result. The modified model is described by the equations:

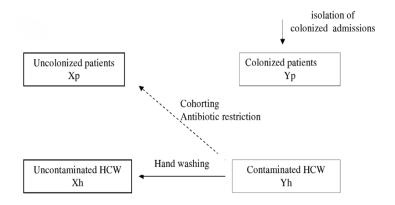


Figure 2.2: Scheme of CPKP diffusion with control mechanism from the article.

$$\frac{dX_P(t)}{dt} = (1 - \varphi)\lambda(B - X_P - Y_P) - \mu_U X_P - \alpha b_P X_P Y_H$$

$$\frac{dY_P(t)}{dt} = \varphi \lambda(B - X_P - Y_P) - \mu_C Y_P + \alpha b_P X_P Y_H$$

$$\frac{dX_H(t)}{dt} = \mu_H Y_H - \alpha b_H Y_P X_H - \alpha_H b_{HH} X_H Y_H$$

$$\frac{dY_H(t)}{dt} = -\mu_H Y_H + \alpha b_H Y_P X_H + \alpha_H b_{HH} X_H Y_H$$
(2.1)

Where X_P is the number of uncolonized patient, Y_P is the number of colonized patient, X_H is the number of uncolonized HCW, Y_H is the number of colonized HCW, λ is the probability to add a new patient, φ is the probability that the added patient is colonized, B is the number of bed, μ_U is the discharge rate for the uncolonized patient, μ_C is the discharge rate for colonized patient, μ_H is the probability of a colonized HCW to become uncolonized, α is the average connectivity of the patient-HCW interaction, b_P is the probability that a colonized patient become colonized interacting with a colonized HCW, b_H is the probability that a colonized HCW become colonized interacting with a colonized patient, b_{HH} is the probability that a colonized HCW become colonized interacting with a colonized HCW and α_H is the average connectivity of the HCW-HCW interaction. In the presence of hand disinfection, the probability of contamination from a patient to an HCW and between HCW's is reduced by p%, where p denotes the hand hygiene compliance rate in the surgical unit. The new model will be:

$$\frac{dX_{P}(t)}{dt} = (1 - \varphi)\lambda(B - X_{P} - Y_{P}) - \mu_{U}X_{P} - \alpha b_{P}X_{P}Y_{H}
\frac{dY_{P}(t)}{dt} = \varphi\lambda(B - X_{P} - Y_{P}) - \mu_{C}Y_{P} + \alpha b_{P}X_{P}Y_{H}
\frac{dX_{H}(t)}{dt} = \mu_{H}Y_{H} - \alpha b_{H}(1 - p)Y_{P}X_{H} - \alpha_{H}b_{HH}(1 - p)X_{H}Y_{H}
\frac{dY_{H}(t)}{dt} = -\mu_{H}Y_{H} + \alpha b_{H}(1 - p)Y_{P}X_{H} + \alpha_{H}b_{HH}(1 - p)X_{H}Y_{H}$$
(2.2)

Another feature that can be taken into account is the effect of antibiotics on colonization. Antibiotics may provide CPKP with a selective growth advantage that will result in higher probability of colonization. Thus, the probability b_P of a patient being colonized

per contact with a contaminated HCW can be decomposed into the product of a baseline probability b_{P0} and a factor f representing the impact of antibiotics. If r is the relative risk of colonization associated with these agents and patients receive them for a fraction d of their stay in the unit, then f = 1 + d(r - 1). Antibiotic restriction policies could target to reducing the duration of administration of these agents from d to d'.

2.2 Mean field results

Solving the system of differential equations 2.2 the results are shown by the figure 2.3a with the critical value of p to avoid the contagion, the initial conditions where a pure uncolonized surgical unit that evolves in time.

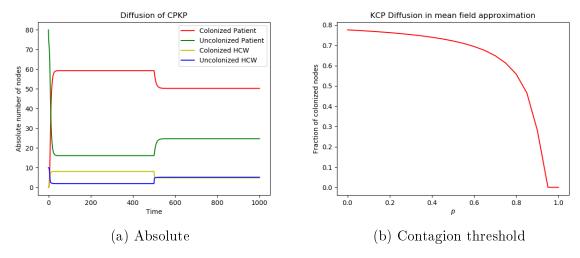


Figure 2.3: Plots of the mean field results.

The figure 2.3a show the absolute number of patients and HCWs in the hypothesis that the control activities are used from the half of the observation time. These results are obtained by solving the differential equation system using a Runge-Kutta at the fourth order in the script $CPKP_Mean_field.py$ and $CPKP_Mean_field_Analysis.py$. As we can see the shape of the relative number of contaminated people as function of the probability of contagion follow the shape of a classical order parameter. The threshold for this model is very high (it depends on the parameters of the model) it is around the value of 1.

In the figure 2.3a we can see that the diffusion of the CPKP and the action of the control activities are very fast this is caused by the high values of the parameter but the value of p(=0.7) is not enough to avoid the contagion because, as it is shown by the Figure 2.3b, the value is below the threshold.

2.3 Application on the network

2.3.1 Different models for the network

In order to implement the model on the network I first had to decide a right network to model the surgical unit. First I decided to consider a big cluster network to model the HCW's interactions and the to add a limited number of patient for each HCW node, than I decide to try to consider a network in which the HCWs shift between the rooms, one time for each time step, in order to interact at least with all the possible patients, in this case the network between the HCWS is absent to simplify the simulation and to have a network which respected the original idea of the model and to reproduce maintenance crew of the HCWs in the surgical unit.

In all the networks I used 10 HCWs and for each of them 8 patients. The network are shown in the Figure 2.4:

- (a) the HCW's cluster has a Barabasi-Albert structure with number of edges to attach from a new node to existing nodes equal to 4
- (b) the Moving-HCW structure with no network between the HCWs
- (c) the HCW's cluster has an Erdos-Renyi structure with probability of having a node 0.5.

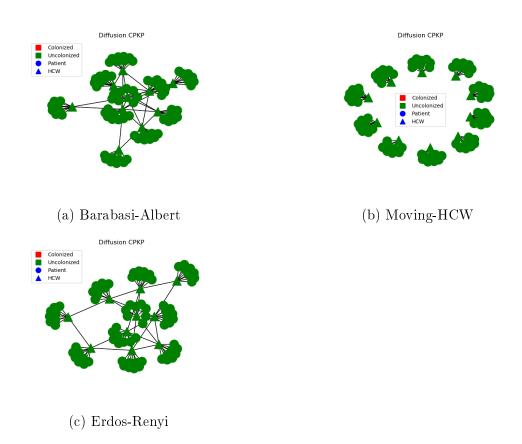
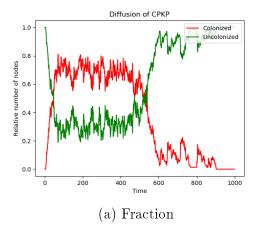


Figure 2.4: Plots of the three networks to model the surgical unit.

2.3.2 Results on a network

The figure 2.5 shows the evolution of the diffusion respect to time, in the figure 2.5a we can see the evolution of the uncolonized and colonized fractions of nodes while in the figure 2.5b we have the evolution of patient and HCW's number. In this simulation the control of the contagion starts at half of the time interval of the observation. The two plots are the results of the code in appendix B. As we can see comparing these results



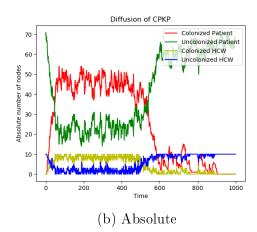


Figure 2.5: Plots of the evolution in time of the diffusion (the network of this simulation is the one with the Erdos-Renyi cluster) using the same parameters and initial conditions of the mean field simulation.

with the mean field one, shown by Figure 2.3a, the epidemic phase of the diffusion reach the same value for the number of colonized and uncolonized patients and HCWs, up to statistical fluctuations, both for network and mean field simulation. The differences comes when we introduce the control activities at half of the simulation time getting a different result because with the same hand hygiene rate in the surgical unit we get a less effect for the mean field result respect to the network. This could be caused by the particular structure of the network which is made by a cluster of HCWs that dominate the diffusion while the patients are only peripheral nodes, an effect is also given by the low number of nodes (90) which doesn't allow the mean field approximation.

2.3.3 Analysis of the effects of the topology on the threshold

In figure 2.6 are shown the plots of the epidemic threshold for the three different networks models.

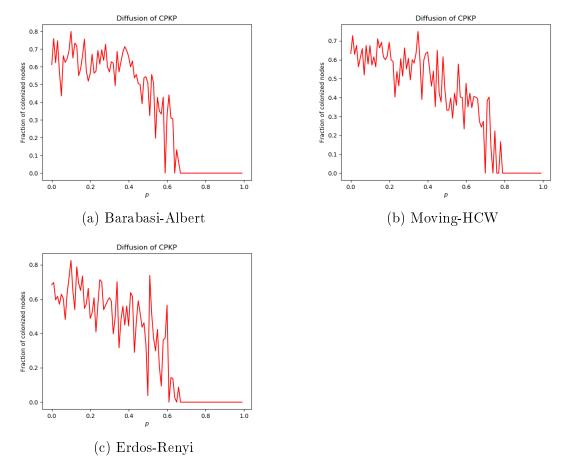


Figure 2.6: Plots of the order parameter in the phase transition between non epidemic and epidemic phase.

As we can see the Barabasi-Albert and the Erdos-Renyi have the same threshold while for the moving HCWs the threshold is bigger, this corroborates the hypothesis of the centrality of the HCW's cluster for the network diffusion. Comparing these plots with the figure 2.3b we see the differences between the network and the mean field diffusion. The first difference is given by the fact that the mean field has a smooth distribution while the network is discontinued but these are statistical fluctuation due to the low number of nodes in the network which doesn't allow the mean field limit. The second and more important difference is the threshold which is bigger for the mean field respect to the network, this can be caused by the structure of the network that, as we have see comparing the different topologies, has a central role in the dynamic of the diffusion. In conclusion in this work we have corroborate the centrality of the topology in order to avoid the contagion in fact as we have seen a fixed room HCWs network is more efficient than a moving HCWs network even if there are contacts between the HCWs, for small number we have seen that the structure of this contact is less relevant than what could be expected.

Appendix A

Python code for CPKP mean field

```
import matplotlib
2 from matplotlib import pyplot as plt
3 import math
4 import random
5 from numpy import *
7 def Deriv(x,y):
      #Parameters of the model
      mu_c=0.05  #Colonized discharge rate
9
     mu_u=0.1  #Uncolonized discharge rate
alpha=0.21  #Probability of colonization
phi=0.05  #Probability that the patient added is colonized
lambd=1  #Probability to add a patient
12
      mu_HC=0.5  #Probability of a colonized HCW to become uncolonized
      p = 0
15
      b = 0.1
      c=0.5
      B = 80
1.8
      dy=zeros(n,'double')
19
2.0
    dy[0] = (1-phi)*lambd*(B-y[0]-y[1])-mu_u*y[0]-alpha*b*y[0]*y[3]
21
      dy[1] = phi*lambd*(B-y[0]-y[1])-mu_c*y[1]+alpha*b*y[0]*y[3]
      dy[2] = mu_HC*y[3] - alpha*b*y[1]*y[2] - alpha*c*y[2]*y[3]
23
      dy[3] = -mu_HC*y[3] + alpha*b*y[1]*y[2] + alpha*c*y[2]*y[3]
      return dy
27 def Derivmod(x,y):
      #Parameters of the model
      mu_c=0.05 #Colonized discharge rate
                      #Uncolonized discharge rate
      mu_u=0.1
30
                     #Probability of colonization
      alpha=0.21
31
      phi=0 #Probability that the patient added is colonized
      lambd=1
                     #Probability to add a patient
      mu_HC=0.5
                     #Probability of a colonized HCW to become uncolonized
34
      p=0.7
      b = 0.1
      c=0.5
      B = 80
38
      dy=zeros(n,'double')
39
```

```
dy[0] = (1-phi)*lambd*(B-y[0]-y[1])-mu_u*y[0]-alpha*b*y[0]*y[3]
41
       dy[1] = phi * lambd*(B-y[0] - y[1]) - mu_c*y[1] + alpha*b*y[0]*y[3]
       dy[2]=mu_HC*y[3]-(1-p)*alpha*b*y[1]*y[2]-(1-p)*alpha*c*y[2]*y[3]
43
       dy[3] = -mu_HC*y[3] + (1-p)*alpha*b*y[1]*y[2] + (1-p)*alpha*c*y[2]*y[3]
44
       return dy
46
  def RK4(x,y,n,h,xf):
47
       k1=zeros(n, 'double')
48
       k2=zeros(n, 'double')
49
                     'double')
       k3=zeros(n,
50
                     'double')
       k4=zeros(n,
51
       ym=zeros(n, 'double')
52
       ye=zeros(n, 'double')
       slope=zeros(n, 'double')
54
       if x < xf/2:
           k1 = Deriv(x,y)
       else:
58
           k1 = Derivmod(x,y)
59
       for i in range(n):
           ym[i] = y[i] + k1[i] * h/2
63
       if x < xf/2:
64
            k2 = Deriv(x+h/2,ym)
66
            k2 = Derivmod(x+h/2,ym)
67
68
       for i in range(n):
69
            ym[i] = y[i] + k2[i] * h/2
       if x < xf/2:
            k3 = Deriv(x+h/2,ym)
73
       else:
74
           k3 = Derivmod(x+h/2,ym)
       for i in range(n):
77
            ye[i] = y[i] + k3[i] * h
78
       if x < xf/2:
79
            k4 = Deriv(x+h, ye)
81
            k4 = Derivmod(x+h, ye)
82
83
       for i in range(n):
84
            slope[i]=(k1[i]+2*(k2[i]+k3[i])+k4[i])/6
85
           y[i]=y[i]+slope[i]*h
86
       x = x + h
       return x, y
89
  def integrator(x,y,n,h,xend,xf):
90
       while x < x end:
91
           if xend-x<h:</pre>
92
                h = x end - x
93
           x, y = RK4(x, y, n, h, xf)
94
95
       return x ,y
```

```
97 n = 4
98 y=zeros(n,'double')
99 yi=zeros(n,'double')
100 yi[0]=80
101 yi[1]=0
102 yi[2]=10
103 yi[3]=0
104 \text{ xi} = 0
105 dx = 0.01
106 \text{ xf} = 1000
107 xout = 1
108
j = int(dx * xf)
xp=zeros(1001, 'double')
yp=zeros((n,1001), 'double')
113 x = xi
114 \text{ m} = 0
115 \text{ xp}[m] = x
for i in range(n):
      yp[i,m]=yi[i]
       y[i]=yi[i]
118
119
120 while x < xf:</pre>
121
      xend=x+xout
       if xend>xf:
122
            xend = xf
123
     h = dx
124
      x,y=integrator(x,y,n,h,xend,xf)
125
      m = m + 1
126
      xp[m] = x
127
       for i in range(n):
129
           yp[i,m]=y[i]
130
#Plot the total number of colonized nodes for time step
plt.plot(xp,yp[1],'r',label='Colonized Patient')
plt.plot(xp,yp[0],'g',label='Uncolonized Patient')
plt.plot(xp,yp[3],'y',label='Colonized HCW')
plt.plot(xp,yp[2],'blue',label='Uncolonized HCW')
plt.legend(loc='upper right')
plt.xlabel('Time')
plt.ylabel('Absolute number of nodes')
plt.title('Diffusion of CPKP')
140 plt.show()
```

Appendix B

Python code for CPKP

The python code for the network diffusion on an Eros-Renyi is based on an event driven algorithm as shown in the section 1.3

```
import matplotlib.pyplot as plt
 2 import math
 3 import random
 4 import graphviz
 5 from networkx import *
 6 from numpy import *
 8 #Preparing the graph
 9 #Num_HCW HCW nodes, each with 8 patient nodes as maximum number
10 \text{ Num} \text{\_HCW} = 10
11 G=fast_gnp_random_graph(Num_HCW, 0.5, seed=None, directed=False)
12 Role = ['HCW']
set_node_attributes(G, Role, 'Role')
                                   #Setting the layout to fix the position
14 fixed_pos=spring_layout(G)
      of the HCW nodes in the plot
for n in range(len(G.nodes())):
                                                     #Adding the patient nodes, 8
      for
                                                     #each HCW
       for i in range(8):
            new = 9*(n+1)+i+1
18
            G.add_node(new,Role=['Patient'])
            G.add_edge(n,new)
22 State=['Uncolonized']
set_node_attributes(G, State,'State') #Setting all the states to
      uncolonized
24 pos = spring_layout(G,pos=fixed_pos,fixed=fixed_pos.keys())
      of the node with fixed the HCW nodes
26 infected_list=[] #Inizialization of the infected list
27 #Parameters of the model
28 \ \text{mu\_c=0.05} #Colonized discharge rate
                 #Uncolonized discharge rate
29 \text{ mu} \, \underline{\text{u}} = 0.1
alpha=0.21 #Probability of colonization
phi=0.05 #Probability that the patient added is colonized
lambd=1 #Probability to add a patient
mu_HC=0.5 #Probability of a colonized HCW to become uncolonized
                   #Probability of a colonized HCW to become uncolonized
34 \text{ beta} = 0.21
```

```
p = 0.7
36 #Colonized admition rate phi*lambd
37 #Uncolonized admition rate (1-phi)*lambd
38 Time=1000 #Time steps of the simulation (days)
Frac_Col=zeros(Time, 'double')
40 Frac_Uncol=zeros(Time, 'double')
41 Uncol_HCW=zeros(Time, 'double')
42 Col_HCW=zeros(Time, 'double')
43 Uncol_pat=zeros(Time, 'double')
44 Col_pat=zeros(Time, 'double')
45 tim=zeros(Time,'int')
46 #Begin of the time simulation
47 for t in range(Time):
4.8
      #Adding the methods to remove the diffusion after the beginning of
4.9
     the contagion
      if t>Time/2:
          phi=0
                        #Isolation of admitted colonized patient
51
          beta = 0.21*(1-p)
                             #Head washing to decrease the probability to
52
      became colonized after the contact
54
      #Decontaminatio of the HCW
55
      for i in range(Num_HCW):
           if G.nodes[i]['State'] == ['Colonized']:
               rand_head=random.uniform(0,1)
58
               if rand_head <= mu_HC:</pre>
59
                   G.nodes[i]['State']=['Uncolonized']
60
61
    #Removing the nodes
62
      to_remove_list=[]
63
      for node in G.nodes():
           if G.nodes[node]['Role'] == ['Patient']:
65
               if G.nodes[node]['State'] == ['Colonized']:
                   rand_colon=random.uniform(0,1)
67
                   if rand_colon <= mu_c:</pre>
                        to_remove_list.append(node)
69
               if G.nodes[node]['State'] == ['Uncolonized']:
                   rand_uncolon=random.uniform(0,1)
                   if rand_uncolon <= mu_u:</pre>
72
                        to_remove_list.append(node)
73
      for j in range(len(to_remove_list)):
           G.remove_node(to_remove_list[j])
      big_node=amax(G.nodes())+1 #Setting the bigger node to begin the
     adding process
      #Adding the nodes kepping the maximum number of patient for each
     node equal to 8
      to_add_list_col=[]
80
      to_add_list_uncol=[]
81
      to_add_list_link=[]
82
      newnum=big_node
83
      for i in range(Num_HCW):
                                                            #Cicling only on
84
     the HCW because we can only add patient to the HCW
          neighbours=list(G.neighbors(i))
```

```
patient=[j for j in range(len(neighbours)) if G.nodes[
86
      neighbours[j]]['Role']==['Patient']] #Counting the number of patient
       for each node
           patient_num=len(patient)
87
88
           if patient_num<7:</pre>
89
                rand_admis=random.uniform(0,1)
90
                if rand_admis <= lambd:</pre>
91
                    rand_pat=random.uniform(0,1)
                    if rand_pat <= phi:</pre>
93
                         to_add_list_col.append(newnum)
                                                             #add colonized
94
      node
                         to_add_list_link.append((i,newnum))
95
                         newnum = newnum + 1
96
                    else:
97
                         to_add_list_uncol.append(newnum)
                                                              #add uncolonized
9.8
      node
                         to_add_list_link.append((i,newnum))
99
                         newnum = newnum + 1
101
       #Adding of the nodes to the graph
       for i in range(len(to_add_list_col)):
           G.add_node(to_add_list_col[i], Role=['Patient'], State=['
104
      Colonized'])
       for i in range(len(to_add_list_uncol)):
106
           G.add_node(to_add_list_uncol[i], Role=['Patient'], State=['
107
      Uncolonized'])
108
       for i in range(len(to_add_list_link)):
           G.add_edge(*to_add_list_link[i])
       #Stop condition if the graph is empty
       if len(G.nodes()) == 0:
113
           break
114
115
       #Infection for each node
       to_infect_list=[]
117
       for node in G.nodes():
118
           if G.nodes[node]['State'] == ['Colonized']:
                neighbours=list(G.neighbors(node))
                for j in range(len(neighbours)):
                    if G.nodes[neighbours[j]]['State'] == ['Uncolonized']:
                         if G.nodes[neighbours[j]]['Role']==['HCW']:
123
                             rand_inf=random.uniform(0,1)
124
                             if rand_inf <= beta:</pre>
                                  to_infect_list.append(neighbours[j])
126
                         else:
                             rand_inf=random.uniform(0,1)
128
                             if rand_inf <= alpha:</pre>
129
                                  to_infect_list.append(neighbours[j])
132
       to_infect_list=list(set(to_infect_list)) #Keeping only the unique
134
      value for each colonized node
```

```
135
      for i in range(len(to_infect_list)):
           G.nodes[to_infect_list[i]]['State']=['Colonized']
           infected_list.append(to_infect_list[i])
138
139
      #Counting the nodes
140
      for node in G.nodes():
141
           if G.nodes[node]['State'] == ['Colonized']:
142
               Frac_Col[t] = Frac_Col[t]+1
               if G.nodes[node]['Role'] == ['Patient']:
144
                   Col_pat[t] = Col_pat[t] + 1
145
               else:
146
                   Col_HCW[t] = Col_HCW[t] + 1
147
           else:
148
               if G.nodes[node]['Role'] == ['Patient']:
149
                   Uncol_pat[t] = Uncol_pat[t] + 1
               else:
                   Uncol_HCW[t] = Uncol_HCW[t] + 1
      Frac_Col[t] = Frac_Col[t] / len (G. nodes())
      Frac_Uncol[t]=1-Frac_Col[t]
      tim[t]=t
157
158 #Plot the ratio of colonized nodes for time step
plt.plot(tim,Frac_Col,'r',label='Colonized')
plt.plot(tim,Frac_Uncol,'g',label='Uncolonized')
plt.legend(loc='upper right')
plt.xlabel('Time')
plt.ylabel('Relative number of nodes')
plt.title('Diffusion of CPKP')
plt.show()
166 #Plot the total number of colonized nodes for time step
plt.plot(tim,Col_pat,'r',label='Colonized Patient')
plt.plot(tim,Uncol_pat,'g',label='Uncolonized Patient')
plt.plot(tim,Col_HCW,'y',label='Colonized HCW')
plt.plot(tim,Uncol_HCW,'blue',label='Uncolonized HCW')
plt.legend(loc='upper right')
plt.xlabel('Time')
plt.ylabel('Absolute number of nodes')
plt.title('Diffusion of CPKP')
175 plt.show()
```

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