SMA Stable Point Analyses

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1 Endemic Equilibria Stability Analyses of the Exclusive Infection Model from Spicknall et al (2013)

1.1 1.1 Deriving Equilibria and Determining Their Stability

```
[4]: Eqn_S = -beta_W/N*I_W*S-beta_Z/

N*I_Z*S+gamma*(1-epsilon)*I_W+gamma_T*epsilon*I_W+gamma*I_Z

Eqn_I_W = beta_W/N*I_W*S-gamma*(1-epsilon)*I_W-gamma_T*epsilon*I_W

Eqn_I_Z = beta_Z/N*I_Z*S-gamma*I_Z
```

[5]:
$$\begin{bmatrix} I_W \epsilon \gamma_T + I_W \gamma \left(1 - \epsilon\right) - \frac{I_W S \beta_W}{N} + I_Z \gamma - \frac{I_Z S \beta_Z}{N} \\ -I_W \epsilon \gamma_T - I_W \gamma \left(1 - \epsilon\right) + \frac{I_W S \beta_W}{N} \\ -I_Z \gamma + \frac{I_Z S \beta_Z}{N} \end{bmatrix}$$

[6]:
$$\left[(S, 0, 0), \left(\frac{N(-\epsilon \gamma + \epsilon \gamma_T + \gamma)}{\beta_W}, I_W, 0 \right), \left(\frac{N \gamma}{\beta_Z}, 0, I_Z \right) \right]$$

The disease free equilibrium (DFE) is the first vector.

We are not interested in whether the disease free equilibrium (DFE), the first equilibria, is stable. This can be found through deriving R0 using PyGom's matrix methodology for determining R0.

Let's look at the stability of the non-DFEs, following recipe 8.2 of Otto & Day (2007). Needs to create Jacobian Matrix of the models ODEs.

$$\begin{bmatrix} -\frac{I_{W}\beta_{W}}{N} - \frac{I_{Z}\beta_{Z}}{N} & \epsilon\gamma_{T} + \gamma\left(1 - \epsilon\right) - \frac{S\beta_{W}}{N} & \gamma - \frac{S\beta_{Z}}{N} \\ \frac{I_{W}\beta_{W}}{N} & -\epsilon\gamma_{T} - \gamma\left(1 - \epsilon\right) + \frac{S\beta_{W}}{N} & 0 \\ \frac{I_{Z}\beta_{Z}}{N} & 0 & -\gamma + \frac{S\beta_{Z}}{N} \end{bmatrix}$$

Substitute the 1st non-disease free equilibrium (non-DFE) into the Jacobian matrix and determine Eigen values.

[9]:
$$\left[-\frac{\beta_W \gamma + \beta_Z \epsilon \gamma - \beta_Z \epsilon \gamma_T - \beta_Z \gamma}{\beta_W}, -\frac{I_W \beta_W}{N}, 0 \right]$$

Substitute the 2nd non-disease free equilibrium (non-DFE) into the Jacobian matrix and determine Eigen values.

[10]:
$$\left[\frac{\beta_W \gamma + \beta_Z \epsilon \gamma - \beta_Z \epsilon \gamma_T - \beta_Z \gamma}{\beta_Z}, -\frac{I_Z \beta_Z}{N}, 0\right]$$

1.2 Function for Determining the Stable Endemic Equilibrium Infection Prevalence

```
[]: # A rounding to sigfig function will prove useful:
      def round sf(number, significant):
          111
          Rounds to a specified number of significant figures.
          return round(number, significant - len(str(number)))
[12]: def exclus_inf_end_equil(param_values):
          Calculates the non-disease free equilibria for the exclusive infection \Box
          as out lined in Spicknall et al 2013. Returning any non-DFE
          that are biologically reasonable and locally stable.
          # Note if both strains have RO values <1 there is no point in using this,
       \hookrightarrow function.
          equil_pops = []
          # Setup Equilibrium populations as an empty list. If any of the endemic
          \# equilibria are found to biologically feasible and stable they are
       \rightarrow appended to the list.
          #Basic Reproductive numbers from Spicknall 2013
          RO_W = beta_W/(gamma*(1-epsilon)+gamma_T*epsilon)
          RO_W = RO_W.subs(param_values)
          RO_Z = beta_Z/gamma
          RO_Z = RO_Z.subs(param_values)
          #Formula for the non-Disease Free Equilibria has been worked out in a_{\sqcup}
       → jupyter notebook
          #and pasted here. NOTE with the first non-DFE the I W population = N-S_{II}
       \hookrightarrow (I Z=0),
          # and with the second non-DFE the I Z population = N-S (I W=O).
          # Therefore, only the formula for S are needed (see jupyter notebook).
          S_non_DFE_1 = N*(-epsilon*gamma + epsilon*gamma_T + gamma)/beta_W
          S_non_DFE_1 = S_non_DFE_1.subs(param_values)
          I_W_non_DFE_1 = param_values['N']-S_non_DFE_1
          I_Z_{non_DFE_1} = 0
          S non DFE 2 = N*gamma/beta Z
          S_non_DFE_2 = S_non_DFE_2.subs(param_values)
          I_W_non_DFE_2 = 0
          I_Z_non_DFE_2 = param_values['N']-S_non_DFE_2
```

```
#The Eigen values associated with Jacobian matricies for the non-DFE have
\rightarrowbeen
   #derived in a jupyter notebook and pasted here.
   eigs_J_non_DFE_1 = [-(beta_W*gamma + beta_Z*epsilon*gamma -_
→beta_Z*epsilon*gamma_T - beta_Z*gamma)/beta_W,
                        -I_W*beta_W/N,
   eigs_J_non_DFE_2 = [(beta_W*gamma + beta_Z*epsilon*gamma -_
→beta_Z*epsilon*gamma_T - beta_Z*gamma)/beta_Z,
                        -I_Z*beta_Z/N,
   if math.isnan(S_non_DFE_1) or math.isnan(I_W_non_DFE_1) or math.
→isnan(I Z non_DFE 1) or math.isinf(S_non_DFE_1) or math.isinf(I W_non_DFE_1)_
→or math.isinf(I_Z_non_DFE_1):
       DFE_1_feasable = False
   else:
       DFE_1_feasable = True
   if DFE 1 feasable and round sf(S non DFE 1,2) >= 0 and
→round_sf(I_W_non_DFE_1,2) >= 0 and round_sf(I_Z_non_DFE_1,2) >= 0:
       DFE_1_feasable = True
   else:
       DFE_1_feasable = False
   if math.isnan(S non_DFE_2) or math.isnan(I_W_non_DFE_2) or math.
→isnan(I_Z_non_DFE_2) or math.isinf(S_non_DFE_2) or math.isinf(I_W_non_DFE_2)_
→or math.isinf(I_Z_non_DFE_2):
       DFE_2_feasable = False
   else:
       DFE_2_feasable = True
   if DFE_2_feasable and round_sf(S_non_DFE_2,2) >= 0 and_
\rightarrowround_sf(I_W non_DFE_2,2) >= 0 and round_sf(I_Z_non_DFE_2,2) >= 0:
       DFE 2 feasable = True
   else:
       DFE_2_feasable = False
   #Determine if RO_W is >=1, RO_Z<1 and the first DFE is biologically \sqcup
\rightarrow feasible.
   if RO_W.subs(param_values)>=1 and RO_Z.subs(param_values)<1 and_
→DFE_1_feasable:
       equil_pops.append({'S':S_non_DFE_1, 'I_W':I_W_non_DFE_1, 'I_Z':
\rightarrowI Z non DFE 1})
```

```
#Determine if RO Z is \geq1, RO W<1 and the second DFE is biologically.
\hookrightarrow feasible.
   if RO_Z.subs(param_values)>=1 and RO_W.subs(param_values)<1 and_
→DFE 2 feasable:
       equil_pops.append({'S':S_non_DFE_2, 'I_W':I_W_non_DFE_2, 'I_Z':
\hookrightarrow I_Z_non_DFE_2})
   #Determine if RO_W is >=1, RO_Z>=1 and the first DFE is biologically \square
\rightarrow feasible but the second is not.
   if RO_W.subs(param_values)>=1 and RO_Z.subs(param_values)>=1 and_
→DFE_1_feasable and DFE_2_feasable == False:
       equil_pops.append({'S':S_non_DFE_1,'I_W':I_W_non_DFE_1,'I_Z':
\hookrightarrow I_Z_non_DFE_1})
   #Determine if RO W is >=1, RO Z>=1 and the second DFE is biologically.
\rightarrow feasable but the first is not.
   if RO W.subs(param values)>=1 and RO Z.subs(param values)>=1 and
→DFE_1_feasable == False and DFE_2_feasable:
       equil_pops.append({'S':S_non_DFE_2, 'I_W':I_W_non_DFE_2, 'I_Z':
\rightarrowI_Z_non_DFE_2})
    \textit{\#Determine if RO\_W is >=1, RO\_Z>=1 and both DFEs are biologically feasable}. 
   if RO_W.subs(param_values)>=1 and RO_Z.subs(param_values)>=1 and_U
→DFE_1_feasable and DFE_2_feasable:
       # If all the values for non-DFE_1 are >= 0, evaluate eigen values
       # associated with the Jacobian matrix of non-DFE_1.
       # Need to be able to substitute in equilibria formula, as well as the \Box
→param values:
       vals_to_subs = copy.deepcopy(param_values)
       vals to subs['S'] = S non DFE 1
       vals to subs['I W'] = I W non DFE 1
       vals_to_subs['I_Z'] = I_Z_non_DFE_1
       if eigs_J_non_DFE_1[0].subs(vals_to_subs) <=0 and eigs_J_non_DFE_1[1].
⇒subs(vals_to_subs)<=0:
            # If the non-vero eigen values associated with the Jacobian matrix
            # of non-DFE_1 are <= 0, the equil pops is appended with non-DFE_1,
            # as it is locally stable.
            equil_pops.append({'S':S_non_DFE_1, 'I_W':I_W_non_DFE_1, 'I_Z':
\hookrightarrowI_Z_non_DFE_1})
       # If all the values for non-DFE 2 are >= 0, evaluate eigen values
       # associated with the Jacobian matrix of non-DFE_2.
       # Need to be able to substitute in equilibria formula, as well as the
→param values:
       vals_to_subs = copy.deepcopy(param_values)
       vals_to_subs['S'] = S_non_DFE_2
```

```
vals_to_subs['I_W'] = I_W_non_DFE_2
vals_to_subs['I_Z'] = I_Z_non_DFE_2
if eigs_J_non_DFE_2[0].subs(vals_to_subs) <=0 and eigs_J_non_DFE_2[1].

⇒subs(vals_to_subs) <=0:

# If the non-vero eigen values associated with the Jacobian matrix
# of non-DFE_2 are <= 0, the equil_pops is appended with non-DFE_2,
# as it is locally stable.
equil_pops.append({'S':S_non_DFE_2,'I_W':I_W_non_DFE_2,'I_Z':
→I_Z_non_DFE_2})
return(equil_pops)</pre>
```

2 Endemic Equilibria Stability Analyses of the Replacement Infection Model from Spicknall et al (2013)

2.1 2.1 Deriving Equilibria

[14]:
$$\begin{bmatrix} I_W \epsilon \gamma_T + I_W \gamma \left(1 - \epsilon \right) - \frac{I_W S \beta_W}{N} + I_Z \gamma - \frac{I_Z S \beta_Z}{N} \\ I_W I_Z \left(\frac{\beta_W}{N} - \frac{\beta_Z}{N} \right) - I_W \epsilon \gamma_T - I_W \gamma \left(1 - \epsilon \right) + \frac{I_W S \beta_W}{N} \end{bmatrix}$$
$$I_W I_Z \left(-\frac{\beta_W}{N} + \frac{\beta_Z}{N} \right) - I_Z \gamma + \frac{I_Z S \beta_Z}{N}$$

$$\left[(S, 0, 0), \left(\frac{I_W \beta_W - I_W \beta_Z + N \gamma}{\beta_Z}, I_W, \frac{-I_W \beta_W^2 + I_W \beta_W \beta_Z - N \beta_W \gamma - N \beta_Z \epsilon \gamma + N \beta_Z \epsilon \gamma_T + N \beta_Z \gamma}{\beta_Z (\beta_W - \beta_Z)} \right), \left(\frac{N (-1)^2 \beta_W^2 + N \beta_W \beta_W - N \beta_W \gamma -$$

The 2nd vector is difficult to interprate I_W appearing in all three elements, it requires some substitution work.

2.1.1 2.1.1 Dealing with Unusual Equilibria

$$\left(\frac{I_W\beta_W - I_W\beta_Z + N\gamma}{\beta_Z}, \ I_W, \ \frac{-I_W\beta_W^2 + I_W\beta_W\beta_Z - N\beta_W\gamma - N\beta_Z\epsilon\gamma + N\beta_Z\epsilon\gamma_T + N\beta_Z\gamma}{\beta_Z\left(\beta_W - \beta_Z\right)}\right)$$

2.1.1.1 Treating as a Simultaneous Equation Problem.

```
[17]: exp2_as_simul=unusaul_equil[1]-I_W
exp3_as_simul=unusaul_equil[2]-I_Z
sol = sympy.solve((exp2_as_simul,exp3_as_simul),(I_W,I_Z))
sol
```

[17]:
$$\left\{ I_W : -\frac{I_Z \beta_Z}{\beta_W} + \frac{-N\beta_W \gamma - N\beta_Z \epsilon \gamma + N\beta_Z \epsilon \gamma_T + N\beta_Z \gamma}{\beta_W^2 - \beta_W \beta_Z} \right\}$$

2.1.1.2 Substituting $S = N - (I_W + I_Z)$ and treating as a Simultaneous Equation Problem. Lets try substituting $S = N - (I_W + I_Z)$ into the 2nd and 3rd equation:

 I_W

$$\frac{-I_W \beta_W^2 + I_W \beta_W \beta_Z - N \beta_W \gamma - N \beta_Z \epsilon \gamma + N \beta_Z \epsilon \gamma_T + N \beta_Z \gamma}{\beta_Z (\beta_W - \beta_Z)}$$

Solving both equations for I_W and I_Z

[19]:
$$\left\{I_W: -\frac{I_Z\beta_Z}{\beta_W} + \frac{-N\beta_W\gamma - N\beta_Z\epsilon\gamma + N\beta_Z\epsilon\gamma_T + N\beta_Z\gamma}{\beta_W^2 - \beta_W\beta_Z}\right\}$$

There is no solution to this approach!

2.2.1.3 Substituting $N = S + I_W + I_Z$ and treating as a Simultaneous Equation Problem. Lets try substituting $N = S + I_W + I_Z$ into the 2nd and 3rd equation:

 I_{W}

$$\frac{-I_{W}\beta_{W}^{2}+I_{W}\beta_{W}\beta_{Z}-\beta_{W}\gamma\left(I_{W}+I_{Z}+S\right)-\beta_{Z}\epsilon\gamma\left(I_{W}+I_{Z}+S\right)+\beta_{Z}\epsilon\gamma_{T}\left(I_{W}+I_{Z}+S\right)+\beta_{Z}\gamma\left(I_{W}+I_{Z}+S\right)}{\beta_{Z}\left(\beta_{W}-\beta_{Z}\right)}$$

$$\left\{I_W: \frac{I_Z\left(-\beta_W\beta_Z - \beta_W\gamma + \beta_Z^2 - \beta_Z\epsilon\gamma + \beta_Z\epsilon\gamma_T + \beta_Z\gamma\right)}{\beta_W^2 - \beta_W\beta_Z + \beta_W\gamma + \beta_Z\epsilon\gamma - \beta_Z\epsilon\gamma_T - \beta_Z\gamma} + \frac{-S\beta_W\gamma - S\beta_Z\epsilon\gamma + S\beta_Z\epsilon\gamma_T + S\beta_Z\gamma}{\beta_W^2 - \beta_W\beta_Z + \beta_W\gamma + \beta_Z\epsilon\gamma - \beta_Z\epsilon\gamma_T - \beta_Z\gamma}\right\}$$

Whilst not getting rid of the S term in the equations for I_W and I_Z , this does reveal that this equilibrium point is biologically meaningless. Either $\beta_W < \beta_Z$ and I_W is negative, $\beta_Z < \beta_W$ and I_Z is negative or $\beta_W = \beta_Z$ in which case $I_W = undefined$ and $I_Z = -undefined$.

2.2 2.2 Determining Equilibria Stability

We are not interested in whether the disease free equilibrium (DFE), the first equilibria, is stable. This can be found through deriving R0 using PyGom's matrix methodology for determining R0.

Lets look at the stability of the non-DFEs, following recipe 8.2 of Otto & Day (2007). Needs to create Jacobian Matrix of the models ODEs.

The 2nd equilibrium as discussed in 1.2.1 is biologically meaningless. Therefore, we ignore this equilibrium.

$$\begin{bmatrix} -\frac{I_{W}\beta_{W}}{N} - \frac{I_{Z}\beta_{Z}}{N} & \epsilon\gamma_{T} + \gamma\left(1 - \epsilon\right) - \frac{S\beta_{W}}{N} & \gamma - \frac{S\beta_{Z}}{N} \\ \frac{I_{W}\beta_{W}}{N} & I_{Z}\left(\frac{\beta_{W}}{N} - \frac{\beta_{Z}}{N}\right) - \epsilon\gamma_{T} - \gamma\left(1 - \epsilon\right) + \frac{S\beta_{W}}{N} & I_{W}\left(\frac{\beta_{W}}{N} - \frac{\beta_{Z}}{N}\right) \\ \frac{I_{Z}\beta_{Z}}{N} & I_{Z}\left(-\frac{\beta_{W}}{N} + \frac{\beta_{Z}}{N}\right) & I_{W}\left(-\frac{\beta_{W}}{N} + \frac{\beta_{Z}}{N}\right) - \gamma + \frac{S\beta_{Z}}{N} \end{bmatrix}$$

Substitute the 1st non-disease free equilibrium (non-DFE) into the Jacobian matrix and determine Eigen values.

$$\left[-\frac{I_W \beta_W^2 - I_W \beta_W \beta_Z + N \beta_W \gamma + N \beta_Z \epsilon \gamma - N \beta_Z \epsilon \gamma_T - N \beta_Z \gamma}{N \beta_W}, -\frac{I_W \beta_W}{N}, 0 \right]$$

Substitute the 2nd non-disease free equilibrium (non-DFE) into the Jacobian matrix and determine Eigen values.

```
[24]: non_DFE_2 = {'S':equilibria[3][0],'I_W':equilibria[3][1],'I_Z':equilibria[3][2]} 

J_of_ODEs_non_DFE_2 = J_of_ODEs.subs(non_DFE_2)
J_of_ODEs_non_DFE_2

eigs_J_non_DFE_2 = J_of_ODEs_non_DFE_2.eigenvals()

# The above code line enters the Eigen values as keys to a dictionary.

# It would be more useful to have them as elements in a vector.

eigs_J_non_DFE_2 = [k for k in eigs_J_non_DFE_2.keys()]

eigs_J_non_DFE_2

[24]: \left[\frac{I_Z\beta_W\beta_Z - I_Z\beta_Z^2 + N\beta_W\gamma + N\beta_Z\epsilon\gamma - N\beta_Z\epsilon\gamma_T - N\beta_Z\gamma}{N\beta_Z}, -\frac{I_Z\beta_Z}{N}, 0\right]
```

2.3 Function for Determining the Stable Endemic Equilibrium Infection Prevalence

```
[26]: def replace_inf_end_equil(param_values):
           Calculates the non-disease free equilibria for the replacement infection \Box
       \hookrightarrow model,
           as out lined in Spicknall et al 2013. Reuturning any non-DFE that are
           that are biologically reasonable and locally stable.
           # Note if both strains have RO values <1 there is no point in using this \Box
       \hookrightarrow function.
          equil pops = []
           # Setup Equilibrium populations as an empty list. If any of the endemic
           # equilibria are found to be biologically feasible and stable they are
       \rightarrow appended to the list.
           #Basic Reproductive numbers from Spicknall 2013
          RO_W = beta_W/(gamma*(1-epsilon)+gamma_T*epsilon)
          RO_Z = beta_Z/gamma
           #Formula for the non-Disease Free Equilibria has been worked out in a_{f \sqcup}
       \rightarrow jupyter notebook
           #and pasted here. NOTE with the first non-DFE the I_W population = N-S_{\sqcup}
       \hookrightarrow (I Z=0),
           # and with the second non-DFE the I_Z population = N-S (I_W=0).
           # Therefore, only the formule for S are needed (see jupyter notebook).
          S_non_DFE_1 = N*(-epsilon*gamma + epsilon*gamma_T + gamma)/beta_W
          S_non_DFE_1 = S_non_DFE_1.subs(param_values)
          I_W_non_DFE_1 = param_values['N']-S_non_DFE_1
          I_Z_non_DFE_1 = 0
          S_{non_DFE_2} = N*gamma/beta_Z
```

```
S_non_DFE_2 = S_non_DFE_2.subs(param_values)
   I_W_{non_DFE_2} = 0
   I_Z_non_DFE_2 = param_values['N']-S_non_DFE_2
   #The eigen values associated with Jacobian matricies for the non-DFE have
⇒been
   #worked out in a jupyter notebook and pasted here.
   eigs_J_non_DFE_1 = [-I_W*beta_W/N,
                        -(I_W*beta_W**2 - I_W*beta_W*beta_Z + N*beta_W*gamma +
→N*beta Z*epsilon*gamma - N*beta Z*epsilon*gamma T - N*beta Z*gamma)/
\hookrightarrow (N*beta_W),
                        0]
   eigs_J_non_DFE_2 = [-I_Z*beta_Z/N,
                        (I_Z*beta_W*beta_Z - I_Z*beta_Z**2 + N*beta_W*gamma +_
→N*beta_Z*epsilon*gamma - N*beta_Z*epsilon*gamma_T - N*beta_Z*gamma)/
\hookrightarrow (N*beta_Z),
                        0]
   if math.isnan(S_non_DFE_1) or math.isnan(I_W_non_DFE_1) or math.
\rightarrow isnan(I_Z_non_DFE_1) or math.isinf(S_non_DFE_1) or math.isinf(I_W_non_DFE_1)_\sqcup
→or math.isinf(I_Z_non_DFE_1):
       DFE 1 feasable = False
   else:
       DFE_1_feasable = True
   if DFE_1_feasable and round_sf(S_non_DFE_1,2) >= 0 and_
→round_sf(I_W_non_DFE_1,2) >= 0 and round_sf(I_Z_non_DFE_1,2) >= 0:
       DFE 1 feasable = True
   else:
       DFE_1_feasable = False
   if math.isnan(S_non_DFE_2) or math.isnan(I_W_non_DFE_2) or math.
→isnan(I_Z_non_DFE_2) or math.isinf(S_non_DFE_2) or math.isinf(I_W_non_DFE_2)_
→or math.isinf(I_Z_non_DFE_2):
       DFE_2_feasable = False
   else:
       DFE_2_feasable = True
   if DFE_2_feasable and round_sf(S_non_DFE_2,2) >= 0 and_
\rightarrowround_sf(I_W_non_DFE_2,2) >= 0 and round_sf(I_Z_non_DFE_2,2) >= 0:
       DFE_2_feasable = True
   else:
       DFE_2_feasable = False
   #Determine if RO W is >=1, RO Z<1 and the first DFE is biologically.
\rightarrow feasable.
```

```
if RO_W.subs(param_values)>=1 and RO_Z.subs(param_values)<1 and__
→DFE_1_feasable:
       equil_pops.append({'S':S_non_DFE_1, 'I_W':I_W_non_DFE_1, 'I_Z':
\rightarrowI Z non DFE 1})
   #Determine if RO Z is \geq1, RO W<1 and the second DFE is biologically.
\hookrightarrow feasable.
   if RO Z.subs(param values)>=1 and RO W.subs(param values)<1 and
→DFE 2 feasable:
       equil_pops.append({'S':S_non_DFE_2, 'I_W':I_W_non_DFE_2, 'I_Z':
\hookrightarrow I_Z_non_DFE_2})
   #Determine if RO W is >=1, RO Z>=1 and the first DFE is biologically,
\rightarrow feasable but the second is not.
   if RO W.subs(param_values)>=1 and RO_Z.subs(param_values)>=1 and_
→DFE_1_feasable and DFE_2_feasable == False:
       equil_pops.append({'S':S_non_DFE_1, 'I_W':I_W_non_DFE_1, 'I_Z':
\hookrightarrow I_Z_non_DFE_1})
   #Determine if RO_W is \geq=1, RO_Z\geq=1 and the second DFE is biologically_\square
\rightarrow feasable but the first is not.
   if RO W.subs(param values)>=1 and RO Z.subs(param values)>=1 and
→DFE_1_feasable == False and DFE_2_feasable:
       equil_pops.append({'S':S_non_DFE_2, 'I_W':I_W_non_DFE_2, 'I_Z':
\hookrightarrowI_Z_non_DFE_2})
   \#Determine if RO\ W is >=1, RO\ Z>=1 and both DFEs are biologically feasable.
   if RO W.subs(param values)>=1 and RO Z.subs(param values)>=1 and
→DFE_1_feasable and DFE_2_feasable:
       # If all the values for non-DFE_1 are >= 0, evaluate eigen values
       # associated with the Jacobian matrix of non-DFE_1.
       # Need to be able to substitute in equilibria formula, as well as the
→param values:
       vals to subs = copy.deepcopy(param values)
       vals_to_subs['S'] = S_non_DFE_1
       vals_to_subs['I_W'] = I_W_non_DFE_1
       vals_to_subs['I_Z'] = I_Z_non_DFE_1
       if eigs_J_non_DFE_1[0].subs(vals_to_subs) <=0 and eigs_J_non_DFE_1[1].
⇒subs(vals_to_subs) <=0:
            # If the non-vero eigen values associated with the Jacobian matrix
            # of non-DFE_1 are <= 0, the equil_pops is appended with non-DFE_1,
           # as it is locally stable.
           equil_pops.append({'S':S_non_DFE_1, 'I_W':I_W_non_DFE_1, 'I_Z':
\rightarrowI Z non DFE 1})
       # If all the values for non-DFE 2 are >= 0, evaluate eigenvalues
```

- 3 Endemic Equilibria Stability Analyses of the Bi-Conversion Model from Spicknall et al (2013)
- 3.1 3.1 Deriving Equilibria & The Eigen Values Associated with Their Jacobian Matricies

```
\left(\frac{N\left(-\beta_{W}\epsilon\phi+\beta_{W}\gamma+\beta_{W}\phi-\beta_{Z}\epsilon\gamma+\beta_{Z}\epsilon\gamma_{T}+\beta_{Z}\epsilon\rho+\beta_{Z}\gamma+\sqrt{\beta_{W}^{2}\epsilon^{2}\phi^{2}-2\beta_{W}^{2}\epsilon\gamma\phi-2\beta_{W}^{2}\epsilon\phi^{2}+\beta_{W}^{2}\gamma^{2}+2\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_{W}^{2}\gamma\phi+\beta_
```

The disease free equilibrium (DFE) is the first vector.

[31]: DFE = equilibria[0]

We are not interested in whether the disease free equilibrium (DFE), the first equilibria, is stable. This can be found through deriving R0 using PyGom's matrix methodology for determining R0.

Lets look at the stability of the non-DFEs, following recipe 8.2 of Otto & Day (2007). Needs to create Jacobian Matrix of the models ODEs.

$$\begin{bmatrix} -\frac{I_{W}\beta_{W}}{N} - \frac{I_{Z}\beta_{Z}}{N} & \epsilon\gamma_{T} + \gamma\left(1 - \epsilon\right) - \frac{S\beta_{W}}{N} & \gamma - \frac{S\beta_{Z}}{N} \\ \frac{I_{W}\beta_{W}}{N} & -\epsilon\gamma_{T} - \epsilon\rho - \gamma\left(1 - \epsilon\right) + \frac{S\beta_{W}}{N} & \phi\left(1 - \epsilon\right) \\ \frac{I_{Z}\beta_{Z}}{N} & \epsilon\rho & -\gamma - \phi\left(1 - \epsilon\right) + \frac{S\beta_{Z}}{N} \end{bmatrix}$$

Substitute the 1st non-disease free equilibrium (non-DFE) into the Jacobian matrix and determine Eigen values.

[34]: sympy.simplify(eigs_J_non_DFE_1[0])

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packages\IPython\lib\latextools.py:126: MatplotlibDeprecationWarning:

The to_png function was deprecated in Matplotlib 3.4 and will be removed two minor releases later. Use mathtext.math_to_image instead.

mt.to_png(f, s, fontsize=12, dpi=dpi, color=color)

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packages\IPython\lib\latextools.py:126: MatplotlibDeprecationWarning:

The to_rgba function was deprecated in Matplotlib 3.4 and will be removed two minor releases later. Use mathtext.math_to_image instead.

mt.to_png(f, s, fontsize=12, dpi=dpi, color=color)

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packages\IPython\lib\latextools.py:126: MatplotlibDeprecationWarning:

The to_mask function was deprecated in Matplotlib 3.4 and will be removed two minor releases later. Use mathtext.math_to_image instead.

```
mt.to_png(f, s, fontsize=12, dpi=dpi, color=color)
                                  C:\Users\mdgru\anaconda3\envs\amr\lib\site-
                                  packages\IPython\lib\latextools.py:126: MatplotlibDeprecationWarning:
                                  The MathtextBackendBitmap class was deprecated in Matplotlib 3.4 and will be
                                  removed two minor releases later. Use mathtext.math to image instead.
                                             mt.to_png(f, s, fontsize=12, dpi=dpi, color=color)
[34]:
                                   -I_{Z}\beta_{W}\beta_{Z}^{2}\epsilon\rho+I_{Z}\beta_{W}\beta_{Z}\epsilon\left(\beta_{W}\phi-\beta_{Z}\gamma+\beta_{Z}\gamma_{T}\right)+I_{Z}\beta_{W}\beta_{Z}\left(-\beta_{W}\gamma-\beta_{W}\phi+\beta_{Z}\gamma-\sqrt{\beta_{W}^{2}\epsilon^{2}\phi^{2}-2\beta_{W}^{2}\epsilon\gamma\phi-2\beta_{W}^{2}\epsilon\phi\phi}\right)
[35]: sympy.simplify(eigs_J_non_DFE_1[1])
 [35]:
                                  -I_{Z}\beta_{W}\beta_{Z}^{2}\epsilon\rho + I_{Z}\beta_{W}\beta_{Z}\epsilon\left(\beta_{W}\phi - \beta_{Z}\gamma + \beta_{Z}\gamma_{T}\right) + I_{Z}\beta_{W}\beta_{Z}\left(-\beta_{W}\gamma - \beta_{W}\phi + \beta_{Z}\gamma - \sqrt{\beta_{W}^{2}\epsilon^{2}\phi^{2} - 2\beta_{W}^{2}\epsilon\gamma\phi - 2\beta_{W}^{2}\epsilon\gamma\phi} - 2\beta_{W}^{2}\epsilon\gamma\phi - 2\beta_{W}^{2}\epsilon\gamma\phi\right) + I_{Z}\beta_{W}\beta_{Z}^{2}\epsilon\rho + I_{Z}\beta_{W}\beta_{Z}\epsilon\left(\beta_{W}\phi - \beta_{Z}\gamma + \beta_{Z}\gamma_{T}\right) + I_{Z}\beta_{W}\beta_{Z}\left(-\beta_{W}\gamma - \beta_{W}\phi + \beta_{Z}\gamma - \sqrt{\beta_{W}^{2}\epsilon^{2}\phi^{2} - 2\beta_{W}^{2}\epsilon\gamma\phi - 2\beta_{W}^{2}\epsilon\gamma\phi} - 2\beta_{W}^{2}\epsilon\gamma\phi\right) + I_{Z}\beta_{W}\beta_{Z}^{2}\epsilon\rho + I_{Z}
[36]: sympy.simplify(eigs_J_non_DFE_1[2])
[36]: 0
                                  Substitute the 2nd non-disease free equilibrium (non-DFE) into the Jacobian matrix and determine
                                  Eigen values.
[37]: non_DFE_2 = {'S':equilibria[2][0], 'I_W':equilibria[2][1], 'I_Z':equilibria[2][2]}
                                      J_of_ODEs_non_DFE_2 = J_of_ODEs.subs(non_DFE_2)
                                      J_of_ODEs_non_DFE_2
                                      eigs_J_non_DFE_2 = J_of_ODEs_non_DFE_2.eigenvals()
                                      # The above code line enters the Eigen values as keys to a dictionary.
                                      # It would be more useful to have them as elements in a vector.
                                      eigs_J_non_DFE_2 = [k for k in eigs_J_non_DFE_2.keys()]
[38]: sympy.simplify(eigs_J_non_DFE_2[0])
[38]:
                                  -I_{Z}\beta_{W}\beta_{Z}^{2}\epsilon\rho+I_{Z}\beta_{W}\beta_{Z}\epsilon\left(\beta_{W}\phi-\beta_{Z}\gamma+\beta_{Z}\gamma_{T}\right)+I_{Z}\beta_{W}\beta_{Z}\left(-\beta_{W}\gamma-\beta_{W}\phi+\beta_{Z}\gamma+\sqrt{\beta_{W}^{2}\epsilon^{2}\phi^{2}-2\beta_{W}^{2}\epsilon\gamma\phi-2\beta_{W}^{2}\epsilon\phi\phi}\right)
[39]: sympy.simplify(eigs_J_non_DFE_2[1])
[39]:
                                  -I_{Z}\beta_{W}\beta_{Z}^{2}\epsilon\rho + I_{Z}\beta_{W}\beta_{Z}\epsilon\left(\beta_{W}\phi - \beta_{Z}\gamma + \beta_{Z}\gamma_{T}\right) + I_{Z}\beta_{W}\beta_{Z}\left(-\beta_{W}\gamma - \beta_{W}\phi + \beta_{Z}\gamma + \sqrt{\beta_{W}^{2}\epsilon^{2}\phi^{2} - 2\beta_{W}^{2}\epsilon\gamma\phi - 2\beta_{W}^{2}\epsilon\phi\phi}\right) + I_{Z}\beta_{W}\beta_{Z}^{2}\epsilon\rho + I_{Z}\beta_{W}\beta_{Z}\epsilon\left(\beta_{W}\phi - \beta_{Z}\gamma + \beta_{Z}\gamma_{T}\right) + I_{Z}\beta_{W}\beta_{Z}\left(-\beta_{W}\gamma - \beta_{W}\phi + \beta_{Z}\gamma + \sqrt{\beta_{W}^{2}\epsilon^{2}\phi^{2} - 2\beta_{W}^{2}\epsilon\gamma\phi - 2\beta_{W}^{2}\epsilon\phi\phi}\right) + I_{Z}\beta_{W}\beta_{Z}^{2}\epsilon\rho + I_{Z}
[40]: sympy.simplify(eigs_J_non_DFE_2[2])
[40]: 0
```

3.2 Function for Determining the Stable Endemic Equilibrium Infection Prevelance

```
[42]: def bi_directional_end_equil(param_values):
          Calculates the non-disease free equilibria for the uni and bi-directional \sqcup
       \hookrightarrow convertion model ,
          as out lined in Spicknall et al 2013. Returning any non-DFE that are
          biologically reasonable and locally stable.
          \# Note if both strains have RO values <1 there is no point in using this \sqcup
       \hookrightarrow function.
          equil_pops = []
          # Setup Equilibrium populations as an empty list. If any of the endemic
          # equilibria are found to be biologically feasible they are appended to the
       \hookrightarrow list.
          #Basic Reproductive numbers from Spicknall 2013
          RO_W = 2*beta_W*beta_Z/
       → (beta_W*gamma+beta_Z*gamma-beta_Z*epsilon*gamma+beta_Z*epsilon*gamma_T+beta_W*phi-beta_W*ep
       →((beta_W*(gamma+phi-epsilon*phi)+beta_Z*(gamma-epsilon*gamma+epsilon*(gamma_T+rho)))**2+4*b
       → (-1+epsilon)*epsilon*gamma_T*phi-gamma*(phi+phi*epsilon**2+epsilon*(gamma_T-2*phi+rho))))**
       →5)
          RO_Z = 2*beta_W*beta_Z/
       → (beta_W*gamma+beta_Z*gamma-beta_Z*epsilon*gamma+beta_Z*epsilon*gamma_T+beta_W*phi-beta_W*ep
       →((beta_W*(gamma+phi-epsilon*phi)+beta_Z*(gamma-epsilon*gamma+epsilon*(gamma_T+rho)))**2+4*b
       → (-1+epsilon)*epsilon*gamma_T*phi-gamma*(phi+phi*epsilon**2+epsilon*(gamma_T-2*phi+rho))))**
          #Formula for the non-Disease Free Equilibria has been worked out in a_{\sqcup}
       \rightarrow jupyter notebook
          #and pasted here.
          non_DFE_1 = {
```

```
'S':N*(-beta_W*epsilon*phi + beta_W*gamma + beta_W*phi -
⇒beta_Z*epsilon*gamma + beta_Z*epsilon*gamma_T + beta_Z*epsilon*rho +
→beta_Z*gamma - sqrt(beta_W**2*epsilon**2*phi**2 -
→2*beta_W**2*epsilon*gamma*phi - 2*beta_W**2*epsilon*phi**2 +
→beta_W**2*gamma**2 + 2*beta_W**2*gamma*phi + beta_W**2*phi**2 -
→2*beta_W*beta_Z*epsilon**2*gamma*phi +
→2*beta W*beta Z*epsilon**2*gamma T*phi - 2*beta W*beta Z*epsilon**2*phi*rho
→+ 2*beta_W*beta_Z*epsilon*gamma**2 - 2*beta_W*beta_Z*epsilon*gamma*gamma_T +
→4*beta W*beta Z*epsilon*gamma*phi - 2*beta W*beta Z*epsilon*gamma*rho -
→2*beta_W*beta_Z*epsilon*gamma_T*phi + 2*beta_W*beta_Z*epsilon*phi*rho -
→2*beta_W*beta_Z*gamma**2 - 2*beta_W*beta_Z*gamma*phi +
→beta_Z**2*epsilon**2*gamma**2 - 2*beta_Z**2*epsilon**2*gamma*gamma_T -
→2*beta_Z**2*epsilon**2*gamma*rho + beta_Z**2*epsilon**2*gamma_T**2 +
→2*beta_Z**2*epsilon**2*gamma_T*rho + beta_Z**2*epsilon**2*rho**2 -
→2*beta Z**2*epsilon*gamma**2 + 2*beta Z**2*epsilon*gamma*gamma T +
→2*beta_Z**2*epsilon*gamma*rho + beta_Z**2*gamma**2))/(2*beta_W*beta_Z)
           'I_W':-I_Z*(beta_W*epsilon*phi - beta_W*gamma - beta_W*phi -
→beta Z*epsilon*gamma + beta Z*epsilon*gamma T + beta Z*epsilon*rho +
→beta_Z*gamma - sqrt(beta_W**2*epsilon**2*phi**2 -
→2*beta W**2*epsilon*gamma*phi - 2*beta W**2*epsilon*phi**2 +
→beta_W**2*gamma**2 + 2*beta_W**2*gamma*phi + beta_W**2*phi**2 -□
→2*beta_W*beta_Z*epsilon**2*gamma*phi +
→2*beta W*beta Z*epsilon**2*gamma T*phi - 2*beta W*beta Z*epsilon**2*phi*rho⊔
→+ 2*beta_W*beta_Z*epsilon*gamma**2 - 2*beta_W*beta_Z*epsilon*gamma*gamma_T +
→4*beta_W*beta_Z*epsilon*gamma*phi - 2*beta_W*beta_Z*epsilon*gamma*rho -
→2*beta_W*beta_Z*epsilon*gamma_T*phi + 2*beta_W*beta_Z*epsilon*phi*rho -
→2*beta_W*beta_Z*gamma**2 - 2*beta_W*beta_Z*gamma*phi +
→beta Z**2*epsilon**2*gamma**2 - 2*beta Z**2*epsilon**2*gamma*gamma T -
→2*beta Z**2*epsilon**2*gamma*rho + beta Z**2*epsilon**2*gamma T**2 +
→2*beta_Z**2*epsilon**2*gamma_T*rho + beta_Z**2*epsilon**2*rho**2 -
→2*beta Z**2*epsilon*gamma**2 + 2*beta Z**2*epsilon*gamma*gamma T + 11
→2*beta_Z**2*epsilon*gamma*rho + beta_Z**2*gamma**2))/(2*beta_W*epsilon*rho)
          'I_Z':I_Z
   # Need to track proportion so a value for I Z=1 needs to substituted along \Box
\rightarrow with params values.
   vals_to_subs = copy.deepcopy(param_values)
   vals to subs['I Z'] = 1
   S non DFE 1 = non DFE 1['S'].subs(param values)
   not_S_non_DFE_1 = param_values['N']-S_non_DFE_1
   prop I W non DFE 1 = non DFE 1['I W'].subs(vals to subs)/

→ (1+non_DFE_1['I_W'].subs(vals_to_subs))
   if math.isnan(prop_I_W_non_DFE_1):
      prop_I_W_non_DFE_1 =1
```

```
I_W_non_DFE_1 = (not_S_non_DFE_1*prop_I_W_non_DFE_1)
  I_Z_non_DFE_1 = (not_S_non_DFE_1*(1-prop_I_W_non_DFE_1))
  non_DFE_2 = {
           'S':N*(-beta_W*epsilon*phi + beta_W*gamma + beta_W*phi -
→beta_Z*epsilon*gamma + beta_Z*epsilon*gamma_T + beta_Z*epsilon*rho +
⇒beta Z*gamma + sqrt(beta W**2*epsilon**2*phi**2 -
→2*beta_W**2*epsilon*gamma*phi - 2*beta_W**2*epsilon*phi**2 +
→beta W**2*gamma**2 + 2*beta W**2*gamma*phi + beta W**2*phi**2 -
→2*beta W*beta Z*epsilon**2*gamma*phi +
→2*beta_W*beta_Z*epsilon**2*gamma_T*phi - 2*beta_W*beta_Z*epsilon**2*phi*rho_
→+ 2*beta_W*beta_Z*epsilon*gamma**2 - 2*beta_W*beta_Z*epsilon*gamma*gamma_T +
→4*beta_W*beta_Z*epsilon*gamma*phi - 2*beta_W*beta_Z*epsilon*gamma*rho -
→2*beta_W*beta_Z*epsilon*gamma_T*phi + 2*beta_W*beta_Z*epsilon*phi*rho -
→2*beta_W*beta_Z*gamma**2 - 2*beta_W*beta_Z*gamma*phi +
→beta Z**2*epsilon**2*gamma**2 - 2*beta Z**2*epsilon**2*gamma*gamma_T -
→2*beta_Z**2*epsilon**2*gamma*rho + beta_Z**2*epsilon**2*gamma_T**2 +
→2*beta_Z**2*epsilon**2*gamma_T*rho + beta_Z**2*epsilon**2*rho**2 -
\hookrightarrow2*beta_Z**2*epsilon*gamma**2 + 2*beta_Z**2*epsilon*gamma*gamma_T +
→2*beta_Z**2*epsilon*gamma*rho + beta_Z**2*gamma**2))/(2*beta_W*beta_Z)
           'I W':-I Z*(beta W*epsilon*phi - beta W*gamma - beta W*phi -
→beta_Z*epsilon*gamma + beta_Z*epsilon*gamma_T + beta_Z*epsilon*rho +
→beta_Z*gamma + sqrt(beta_W**2*epsilon**2*phi**2 -
→2*beta_W**2*epsilon*gamma*phi - 2*beta_W**2*epsilon*phi**2 +
→beta W**2*gamma**2 + 2*beta_W**2*gamma*phi + beta_W**2*phi**2 -□
→2*beta_W*beta_Z*epsilon**2*gamma*phi +
→2*beta_W*beta_Z*epsilon**2*gamma_T*phi - 2*beta_W*beta_Z*epsilon**2*phi*rho_
→+ 2*beta_W*beta_Z*epsilon*gamma**2 - 2*beta_W*beta_Z*epsilon*gamma*gamma_T +__
→4*beta_W*beta_Z*epsilon*gamma*phi - 2*beta_W*beta_Z*epsilon*gamma*rho -
→2*beta_W*beta_Z*epsilon*gamma_T*phi + 2*beta_W*beta_Z*epsilon*phi*rho -
→2*beta W*beta Z*gamma**2 - 2*beta W*beta Z*gamma*phi +
→beta_Z**2*epsilon**2*gamma**2 - 2*beta_Z**2*epsilon**2*gamma*gamma_T -
→2*beta Z**2*epsilon**2*gamma*rho + beta Z**2*epsilon**2*gamma T**2 +
→2*beta_Z**2*epsilon**2*gamma_T*rho + beta_Z**2*epsilon**2*rho**2 -
→2*beta_Z**2*epsilon*gamma**2 + 2*beta_Z**2*epsilon*gamma*gamma_T +
→2*beta_Z**2*epsilon*gamma*rho + beta_Z**2*gamma**2))/(2*beta_W*epsilon*rho)
          'I_Z':I_Z
}
  S_non_DFE_2 = non_DFE_2['S'].subs(param_values)
  not_S_non_DFE_2 = param_values['N']-S_non_DFE_2
  prop_I_W_non_DFE_2 = non_DFE_2['I_W'].subs(vals_to_subs)/
if math.isnan(prop_I_W_non_DFE_2):
      prop_I_W_non_DFE_2 =1
```

```
I_W_non_DFE_2 = (not_S_non_DFE_2*prop_I_W_non_DFE_2)
I_Z_non_DFE_2 = (not_S_non_DFE_2*(1-prop_I_W_non_DFE_2))

#The Eigen values associated with Jacobian matricies for the non-DFE have_□
⇒been
#derived in a jupyter notebook and pasted here.
```

```
0]
   if math.isnan(S non_DFE_1) or math.isnan(I_W_non_DFE_1) or math.
→isnan(I_Z_non_DFE_1) or math.isinf(S_non_DFE_1) or math.isinf(I_W_non_DFE_1)_⊔
→or math.isinf(I_Z_non_DFE_1):
       DFE_1_feasable = False
   else:
       DFE_1_feasable = True
   if DFE_1_feasable and round_sf(S_non_DFE_1,2) >= 0 and_
\rightarrowround_sf(I_W_non_DFE_1,2) >= 0 and round_sf(I_Z_non_DFE_1,2) >= 0:
       DFE 1 feasable = True
   else:
       DFE_1_feasable = False
   if math.isnan(S_non_DFE_2) or math.isnan(I_W_non_DFE_2) or math.
→isnan(I_Z_non_DFE_2) or math.isinf(S_non_DFE_2) or math.isinf(I_W_non_DFE_2)_
→or math.isinf(I_Z_non_DFE_2):
       DFE_2_feasable = False
   else:
       DFE_2_feasable = True
   if DFE_2_feasable and round_sf(S_non_DFE_2,2) >= 0 and_
\rightarrowround_sf(I_W_non_DFE_2,2) >= 0 and round_sf(I_Z_non_DFE_2,2) >= 0:
       DFE_2_feasable = True
   else:
       DFE_2_feasable = False
   #Determine if RO W is \geq=1, RO Z<1 and the first DFE is biologically.
\hookrightarrow feasable.
   if RO_W.subs(param_values)>=1 and RO_Z.subs(param_values)<1 and_
→DFE 1 feasable:
       equil_pops.append({'S':S_non_DFE_1, 'I_W':I_W_non_DFE_1, 'I_Z':
\rightarrowI_Z_non_DFE_1})
   #Determine if RO_Z is >=1, RO_W<1 and the second DFE is biologically.
\rightarrow feasable.
   if RO_Z.subs(param_values)>=1 and RO_W.subs(param_values)<1 and_
→DFE_2_feasable:
       equil_pops.append({'S':S_non_DFE_2, 'I_W':I_W_non_DFE_2, 'I_Z':
\rightarrowI_Z_non_DFE_2})
   #Determine if RO W is \geq 1, RO Z \geq 1 and the first DFE is biologically.
→ feasable but the second is not.
```

```
if RO_W.subs(param_values)>=1 and RO_Z.subs(param_values)>=1 and_U
→DFE_1_feasable and DFE_2_feasable == False:
        equil_pops.append({'S':S_non_DFE_1, 'I_W':I_W_non_DFE_1, 'I_Z':
\rightarrowI Z non DFE 1})
   #Determine if RO W is >=1, RO Z>=1 and the second DFE is biologically.
\rightarrow feasable but the first is not.
   if RO W.subs(param values)>=1 and RO Z.subs(param values)>=1 and II
→DFE_1_feasable == False and DFE_2_feasable:
        equil_pops.append({'S':S_non_DFE_2, 'I_W':I_W_non_DFE_2, 'I_Z':
\hookrightarrowI_Z_non_DFE_2})
   #Determine if RO_W is >=1, RO_Z>=1 and both DFEs are biologically feasable.
   if RO_W.subs(param_values)>=1 and RO_Z.subs(param_values)>=1 and__
→DFE_1_feasable and DFE_2_feasable:
        \# There are several denominators in eigen values associated with both \sqcup
\rightarrowDFEs (see jupyter notebook) that can be 0 if rho or epsilon = 0.
        # This means sympy's substitution function can produce undefined values undefined values.
\hookrightarrow (NaN values).
        # Therefore as Spicknall et al (2013) found that the largest of the RO_{\sqcup}
→values was the RO that described the system.
        # We can choose between the DFE's by using ROs.
       if param values['epsilon']==0 or param values['rho'] == 0:
            if RO_W.subs(param_values)> RO_Z.subs(param_values):
                equil_pops.append({'S':S_non_DFE_1, 'I_W':I_W_non_DFE_1, 'I_Z':
\hookrightarrowI_Z_non_DFE_1})
            else:
                equil_pops.append({'S':S non_DFE_2, 'I_W':I_W non_DFE_2, 'I_Z':
\hookrightarrowI_Z_non_DFE_2})
        else:
            # Need to be able to substitute in equilibria formula, as well as |
\rightarrow the param values:
            vals_to_subs = copy.deepcopy(param_values)
            vals_to_subs['S'] = S_non_DFE_1
            vals_to_subs['I_W'] = I_W_non_DFE_1
            vals_to_subs['I_Z'] = I_Z_non_DFE_1
            # Both DFE associated eigen values can be complex and we are only
            # interested in the real part so:
            eig_1_J_non_DFE_1 = sympy.re(eigs_J_non_DFE_1[0].subs(vals_to_subs))
            eig_2_J_non_DFE_1 = sympy.re(eigs_J_non_DFE_1[1].subs(vals_to_subs))
            if eig_1_J_non_DFE_1 <=0 and eig_2_J_non_DFE_1 <=0:</pre>
                # If the non-vero eigen values associated with the Jacobian
\rightarrow matrix
                # of non-DFE_1 are <= 0, the equil_pops is appended with \square
\rightarrow non-DFE_1,
                # as it is locally stable.
```

```
equil_pops.append({'S':S_non_DFE_1, 'I_W':I_W_non_DFE_1, 'I_Z':
\hookrightarrow I_Z_non_DFE_1})
            # If all the values for non-DFE 2 are >= 0, evaluate eigen values
            # associated with the Jacobian matrix of non-DFE 2.
            # Need to be able to substitute in equilibria formula, as well as |
→ the param values:
           vals_to_subs = copy.deepcopy(param_values)
           vals_to_subs['S'] = S_non_DFE_2
           vals_to_subs['I_W'] = I_W_non_DFE_2
           vals_to_subs['I_Z'] = I_Z_non_DFE_2
            # Both DFE associated eigen values can be complex and we are only
            # interested in the real part so:
           eig_1_J_non_DFE_2 = sympy.re(eigs_J_non_DFE_2[0].subs(vals_to_subs))
           eig_2_J_non_DFE_2 = sympy.re(eigs_J_non_DFE_2[1].subs(vals_to_subs))
           if eig_1_J_non_DFE_2 <=0 and eig_2_J_non_DFE_2 <=0:</pre>
                # If the non-vero eigen values associated with the Jacobian
\rightarrow matrix
                # of non-DFE_2 are <= 0, the equil_pops is appended with_
\rightarrow non-DFE_2,
                # as it is locally stable.
                equil_pops.append({'S':S_non_DFE_2, 'I_W':I_W_non_DFE_2, 'I_Z':
\rightarrowI_Z_non_DFE_2})
   return(equil_pops)
```

4 Endemic Equilibria Stability Analyses of the Super Infection Model from Spicknall et al (2013)

4.1 4.1 Deriving Equilibria

```
\begin{bmatrix} I_{W}\epsilon\gamma_{T} + I_{W}\gamma\left(1-\epsilon\right) + I_{Z}\gamma - \frac{S\beta_{W}(I_{W}+I_{WZ}q)}{N} - \frac{S\beta_{Z}(I_{WZ}q+I_{Z})}{N} \\ -I_{W}\epsilon\gamma_{T} - I_{W}\gamma\left(1-\epsilon\right) - \frac{I_{W}\beta_{Z}(I_{WZ}q+I_{Z})}{N} + I_{WZ}\gamma + \frac{S\beta_{W}(I_{W}+I_{WZ}q)}{N} \\ I_{WZ}\epsilon\gamma_{T} + I_{WZ}\gamma\left(1-\epsilon\right) - I_{Z}\gamma - \frac{I_{Z}\beta_{W}(I_{W}+I_{WZ}q)}{N} + \frac{S\beta_{Z}(I_{WZ}q+I_{Z})}{N} \\ \frac{I_{W}\beta_{Z}(I_{WZ}q+I_{Z})}{N} - I_{WZ}\epsilon\gamma_{T} - I_{WZ}\gamma\left(1-\epsilon\right) - I_{WZ}\gamma + \frac{I_{Z}\beta_{W}(I_{W}+I_{WZ}q)}{N} \end{bmatrix}
```

NOTE NEXT CELL MAY RUN FOREVER. A python script containing the same code as this section was run on a server for a week. After this time no solution was found. Hence the code is silenced.

```
[45]: \#equilibria = sympy.solve(ODE_mat,[S, I_W, I_Z,I_WZ]) \#equilibria
```

5 5. Endemic Equilibria Stability Analyses of the Full Coinfection Model from Spicknall et al (2013)

5.1 5.1 Deriving Equilibria

```
[46]: I_WW, I_ZZ = sympy.symbols('I_WW I_ZZ')
[47]: Eqn_S = -S*beta_W*(I_W + q*(2*I_WW + I_WZ))/N - S*beta_Z*(I_Z + q*(I_WZ + I_WZ)/N - S*beta_Z*(I_Z + I_WZ)/N - S*beta_Z*(I_Z + I_WZ)/N - S*beta_Z*(I_Z + I_WZ + I_WZ)/N - S*beta_Z*(I_Z + I_WZ + 
                                                 →2*I ZZ))/N + gamma*(1-epsilon)*I W + gamma T*epsilon*I W + gamma*I Z +
                                                 →gamma*(1-epsilon)*I_WW + gamma_T*epsilon*I_WW + I_ZZ*gamma
                                          Eqn_I W = S*beta_W*(I W + q*(2*I_WW + I_WZ))/N - I_W*beta_Z*(I_Z + q*(I_WZ + I_WZ))/N - I_W*beta_Z*(I_WZ + I_WZ)/N - I_W*beta_Z*(I_WZ + I_WZ + I_WZ)/N - I_W*beta_Z*(I_WZ + I_WZ + I_W
                                                 \rightarrow2*I_ZZ))/N - I_W*beta_W*(I_W + q*(2*I_WW + I_WZ))/N - gamma*(1-epsilon)*I_W_
                                                →- gamma_T*epsilon*I_W + gamma*I_WZ
                                          Eqn_{I_Z} = S*beta_Z*(I_Z + q*(I_WZ + 2*I_ZZ))/N - I_Z*beta_W*(I_W + q*(2*I_WW + I_WZ))/N - I_Z*beta_W*(I_W + I_WZ)/N - I_Z*beta_W*(I_W + I_
                                                 \rightarrowI_WZ))/N - I_Z*beta_Z*(I_Z + q*(I_WZ + 2*I_ZZ))/N - gamma*I_Z +
                                                →(1-epsilon)*gamma*I_WZ + epsilon*gamma_T*I_WZ
                                          →2*I_ZZ)) - (1-epsilon)*gamma*I_WZ - epsilon*gamma_T*I_WZ - gamma*I_WZ
                                          Eqn_I_WW = I_W*beta_W*(I_W + q*(2*I_WW + I_WZ))/N - gamma*(1-epsilon)*I_WW - __
                                                 ⇒gamma_T*epsilon*I_WW
                                          Eqn_I_ZZ = I_Z*beta_Z*(I_Z + q*(I_WZ + 2*I_ZZ))/N - I_ZZ*gamma
[48]: ODE_mat = sympy.Matrix([Eqn_S,Eqn_I_W,Eqn_I_Z,Eqn_I_WZ,Eqn_I_WW,Eqn_I_ZZ])
                                          ODE_mat
                                         \begin{bmatrix} I_{W}\epsilon\gamma_{T} + I_{W}\gamma\left(1-\epsilon\right) + I_{WW}\epsilon\gamma_{T} + I_{WW}\gamma\left(1-\epsilon\right) + I_{Z}\gamma + I_{ZZ}\gamma - \frac{S\beta_{W}(I_{W} + q(2I_{WW} + I_{WZ}))}{N} - \frac{S\beta_{Z}(I_{Z} + q(I_{WZ} + 2I_{ZZ}))}{N} \\ -I_{W}\epsilon\gamma_{T} - I_{W}\gamma\left(1-\epsilon\right) - \frac{I_{W}\beta_{W}(I_{W} + q(2I_{WW} + I_{WZ}))}{N} - \frac{I_{W}\beta_{Z}(I_{Z} + q(I_{WZ} + 2I_{ZZ}))}{N} + I_{WZ}\gamma + \frac{S\beta_{W}(I_{W} + q(2I_{WW} + I_{WZ}))}{N} \\ I_{WZ}\epsilon\gamma_{T} + I_{WZ}\gamma\left(1-\epsilon\right) - I_{Z}\gamma - \frac{I_{Z}\beta_{W}(I_{W} + q(2I_{WW} + I_{WZ}))}{N} - \frac{I_{Z}\beta_{Z}(I_{Z} + q(I_{WZ} + 2I_{ZZ}))}{N} + \frac{S\beta_{Z}(I_{Z} + q(I_{WZ} + 2I_{ZZ}))}{N} \\ \frac{I_{W}\beta_{Z}(I_{Z} + q(I_{WZ} + 2I_{ZZ}))}{N} - I_{WZ}\epsilon\gamma_{T} - I_{WZ}\gamma\left(1-\epsilon\right) - I_{WZ}\gamma + \frac{I_{Z}\beta_{W}(I_{W} + q(2I_{WW} + I_{WZ}))}{N} \\ \frac{I_{W}\beta_{W}(I_{W} + q(2I_{WW} + I_{WZ}))}{N} - I_{WW}\epsilon\gamma_{T} - I_{WW}\gamma\left(1-\epsilon\right) \\ \frac{I_{Z}\beta_{Z}(I_{Z} + q(I_{WZ} + 2I_{ZZ}))}{N} - I_{ZZ}\gamma \end{bmatrix}
```

NOTE NEXT CELL MAY RUN FOREVER. Finding a solution this way runs into the same problem as the Superinfection model. Hence the code is silenced.

5.2 Endemic Equilibria Stability Analyses of the Full Co-Infection model just Anti-microbial sensitive strain.

[51]:
$$\begin{bmatrix} I_{W}\left(\epsilon\gamma_{T}+\gamma\left(1-\epsilon\right)\right)+I_{WW}\left(\epsilon\gamma_{T}+\gamma\left(1-\epsilon\right)\right)-\frac{S\beta_{W}\left(I_{W}+2I_{WW}q\right)}{N}\\ -I_{W}\left(\epsilon\gamma_{T}+\gamma\left(1-\epsilon\right)\right)-\frac{I_{W}\beta_{W}\left(I_{W}+2I_{WW}q\right)}{N}+\frac{S\beta_{W}\left(I_{W}+2I_{WW}q\right)}{N}\\ \frac{I_{W}\beta_{W}\left(I_{W}+2I_{WW}q\right)}{N}-I_{WW}\left(\epsilon\gamma_{T}+\gamma\left(1-\epsilon\right)\right) \end{bmatrix}$$

$$\left[(S, \ 0, \ 0), \ \left(\frac{-2I_W\beta_Wq + I_W\beta_W - N\epsilon\gamma + N\epsilon\gamma_T + N\gamma}{\beta_W}, \ I_W, \ -\frac{I_W^2\beta_W}{2I_W\beta_Wq + N\epsilon\gamma - N\epsilon\gamma_T - N\gamma} \right) \right]$$

The disease free equilibrium (DFE) is the first vector.

(S, 0, 0)

$$\left(\frac{-2I_W\beta_Wq+I_W\beta_W-N\epsilon\gamma+N\epsilon\gamma_T+N\gamma}{\beta_W},\ I_W,\ -\frac{I_W^2\beta_W}{2I_W\beta_Wq+N\epsilon\gamma-N\epsilon\gamma_T-N\gamma}\right)$$

The endemic equilibrium for the AMS strain is not that useful lets try:

5.2.1 5.1.1 Substituting q = 0.5 in the system.

$$\begin{bmatrix} I_{W}\left(\epsilon\gamma_{T}+\gamma\left(1-\epsilon\right)\right)+I_{WW}\left(\epsilon\gamma_{T}+\gamma\left(1-\epsilon\right)\right)-\frac{S\beta_{W}\left(I_{W}+1.0I_{WW}\right)}{N}\\ -I_{W}\left(\epsilon\gamma_{T}+\gamma\left(1-\epsilon\right)\right)-\frac{I_{W}\beta_{W}\left(I_{W}+1.0I_{WW}\right)}{N}+\frac{S\beta_{W}\left(I_{W}+1.0I_{WW}\right)}{N}\\ \frac{I_{W}\beta_{W}\left(I_{W}+1.0I_{WW}\right)}{N}-I_{WW}\left(\epsilon\gamma_{T}+\gamma\left(1-\epsilon\right)\right) \end{bmatrix} \end{bmatrix}$$

- [55]: equilibria_q_is_a_half = sympy.solve(ODEs_q_is_a_half,[S, I_W, I_WW]) equilibria_q_is_a_half
- $\left[(S, \ 0.0, \ 0.0), \ \left(\frac{N \left(-\epsilon \gamma + \epsilon \gamma_T + \gamma \right)}{\beta_W}, \ I_W, \ \frac{I_W^2 \beta_W}{I_W \beta_W + N \epsilon \gamma N \epsilon \gamma_T N \gamma} \right) \right]$
- [57]: EE_q_is_a_half = equilibria_q_is_a_half[1]
 display(EE_q_is_a_half)

$$\left(\frac{N\left(-\epsilon\gamma+\epsilon\gamma_T+\gamma\right)}{\beta_W},\ I_W,\ -\frac{I_W^2\beta_W}{I_W\beta_W+N\epsilon\gamma-N\epsilon\gamma_T-N\gamma}\right)$$

Solve I_W knowing that $N = S + I_W + I_{WW}$.

- [58]: test_eqn = EE_q_is_a_half[0] + EE_q_is_a_half[1] + EE_q_is_a_half[2] test_eqn
- [58]: $-\frac{I_W^2 \beta_W}{I_W \beta_W + N\epsilon \gamma N\epsilon \gamma_T N\gamma} + I_W + \frac{N\left(-\epsilon \gamma + \epsilon \gamma_T + \gamma\right)}{\beta_W}$
- [59]: test_sol = sympy.solve(test_eqn-N,I_W)
 test_sol
- [59]: $\left[\frac{N\left(-\epsilon\gamma + \epsilon\gamma_T + \gamma\right)\left(\beta_W + \epsilon\gamma \epsilon\gamma_T \gamma\right)}{\beta_W^2}\right]$
- [60]: EE_S_q_is_a_half = EE_q_is_a_half[0]
 EE_I_W_q_is_a_half = test_sol[0]
 display(EE_S_q_is_a_half,EE_I_W_q_is_a_half)
 - $\frac{N\left(-\epsilon\gamma + \epsilon\gamma_T + \gamma\right)}{\beta_W}$
 - $\frac{N\left(-\epsilon\gamma + \epsilon\gamma_T + \gamma\right)\left(\beta_W + \epsilon\gamma \epsilon\gamma_T \gamma\right)}{\beta_W^2}$
- [61]: EE_I_WW_q_is_a_half = sympy.

 →solve(N-(EE_S_q_is_a_half+EE_I_W_q_is_a_half+I_WW),I_WW)

 EE_I_WW_q_is_a_half
- [61]: $\left[\frac{N\left(\beta_W^2 + 2\beta_W\left(\epsilon\gamma \epsilon\gamma_T \gamma\right) + \epsilon^2\gamma^2 2\epsilon^2\gamma\gamma_T + \epsilon^2\gamma_T^2 2\epsilon\gamma^2 + 2\epsilon\gamma\gamma_T + \gamma^2\right)}{\beta_W^2} \right]$
- [62]: EE_I_WW_q_is_a_half = EE_I_WW_q_is_a_half[0].simplify()
 EE_I_WW_q_is_a_half
- [62]: $N\left(\beta_W^2 2\beta_W\left(-\epsilon\gamma + \epsilon\gamma_T + \gamma\right) + \epsilon^2\gamma^2 2\epsilon^2\gamma\gamma_T + \epsilon^2\gamma_T^2 2\epsilon\gamma^2 + 2\epsilon\gamma\gamma_T + \gamma^2\right) \frac{\beta_W^2}{\beta_W^2}$
- [63]: EE_q_is_a_half = sympy.Matrix([EE_S_q_is_a_half, EE_I_W_q_is_a_half, DEE_I_WW_q_is_a_half])
 EE_q_is_a_half

```
\begin{bmatrix} \frac{N(-\epsilon\gamma+\epsilon\gamma_T+\gamma)}{\beta_W} \\ \frac{N(-\epsilon\gamma+\epsilon\gamma_T+\gamma)(\beta_W+\epsilon\gamma-\epsilon\gamma_T-\gamma)}{\beta_W^2} \\ \frac{N(\beta_W^2-2\beta_W(-\epsilon\gamma+\epsilon\gamma_T+\gamma)+\epsilon^2\gamma^2-2\epsilon^2\gamma\gamma_T+\epsilon^2\gamma_T^2-2\epsilon\gamma^2+2\epsilon\gamma\gamma_T+\gamma^2)}{\beta_W^2} \end{bmatrix}
[64]: \begin{bmatrix} \text{EE\_q\_is\_a\_half.subs}(\{'\text{N'}:1\text{e3},'\text{beta\_W'}: 0.04 , '\text{beta\_Z'}: 0.015,'\text{gamma'}: 0. \\ \\ \rightarrow 01,'\text{gamma\_T'}: 0.1,'\text{epsilon'}:0,'\text{q'}:0.5} ) \end{bmatrix}
[64]: \begin{bmatrix} 250.0 \\ 187.5 \\ 562.5 \end{bmatrix}
```