```
In [1]: import numpy as np
        from scipy.integrate import odeint
        from scipy.optimize import curve_fit
        from scipy.optimize import differential evolution
        %matplotlib inline
        import matplotlib.pyplot as plt
        from matplotlib.backends.backend_pdf import PdfPages
        import pandas as pd
In [2]: from endpoint_maker import sqres
        from endpoint maker import func
        from endpoint maker import t
        from endpoint maker import new
        from endpoint maker import lightdata
        from endpoint maker import rawdata
In [3]: if name == ' main ':
            def initparams():
                 \#bounds = ([0.00001,10000],[0,4],[0.00001,1000],[0.00001,1000],[0.00001,100
                 bounds = ([0.001, 10], [0.001, 10], [0.001, 10], [0.001, 10])
                 result = differential evolution(sqres,bounds,maxiter=500,popsize=20,polish=
                 return result
In [4]: initialp = initparams()
        print(initialp)
        message: Optimization terminated successfully.
        success: True
            fun: 12.610919903102086
              x: [ 9.998e+00 1.000e-03 1.000e-03 9.735e+00]
            nit: 96
           nfev: 7760
In [5]: #def tester(t,Kd,n,d2,k2,k3,i):
        def tester(t,d2,k2,k3,k7,i):
            inivalues = [1,0,0,0,0,0,0]
            arrayvalues = np.asarray([])
            #for i in range(len(lightdata[:,0])):
            def I(t):
                tindex = t/5
                 if tindex > 12241:
                    tindex = 12240
                 return lightdata[int(tindex)]
            \#def\ odes(z,t,Kd,n,d2,k2,k3):
            def odes(z,t,d2,k2,k3,k7):
                 Pu, Pb, Pa, mRNA, mCherry1, mCherry2, mCherry3 = z
                 d1 = 0.019905
                k1 = 0.08299
                 Kd = 90.41
                 n = 0.964487
                 #d2= 486.67
```

```
\#k2 = 6.597
    #k3= 0.0539
    d3 = 0.000077
    k4 = 1.25
    d4 = 0.000031
    k5 = 0.00283
    k6 = 0.00283
    Pu = z[0]
    Pb = z[1]
    Pa = z[2]
    mRNA = z[3]
    mCherry1 = z[4]
    mCherry2 = z[5]
    mCherry3 = z[6]
    dPudt = d1*Pb - k1*I(t)**n/(Kd**n+I(t)**n)*Pu
    dPbdt = k1*I(t)**n/(Kd**n+I(t)**n)*Pu - k2*Pb - d1*Pb + d2*Pa
    dPadt = k2*Pb - d2*Pa
    dmRNAdt = k3*Pa - d3*mRNA
    dmCherry1dt = k4*mRNA-(d4 + k5)*mCherry1
    dmCherry2dt = k5*mCherry1-(d4+k6)*mCherry2
    dmCherry3dt = k6*mCherry2 - d4*mCherry3
    return [dPudt,dPbdt,dPadt,dmRNAdt,dmCherry1dt,dmCherry2dt,dmCherry3dt]
\#solver = odeint(odes,inivalues,t,args = (Kd,n,d2,k2,k3),hmax=0.1)
solver = odeint(odes,inivalues,t,args = (d2,k2,k3,k7),hmax=0.1)
mCherryout = solver[:,6]
#mCherryout = mCherryout[0:24480:240]
return mCherryout
```

```
In [6]: print(initialp.x)
```

[9.99813544e+00 1.00048218e-03 1.00001922e-03 9.73461126e+00]

```
In [7]: popt, covt = curve_fit(func,t,new,initialp.x,maxfev=1000000)

#popt, covt = curve_fit(func,t,newdata,initialp.x, maxfev=10000000, bounds=((0.0000 #popt, covt = curve_fit(func,t,newdata,initialp.x, maxfev=1000000)

#Kd,n,d2,k2,k3 = popt
#print('Kd=',Kd,'n=',n,'d2=',d2,'k2=',k2,'k3=',k3)

#d3,k4,d4,k5,k6 = popt
#print('d3=',d3,'k4=',k4,'d4=',d4,'k5=',k5,'k6=',k6)

d2,k2,k3,k7 = popt
print('d2=',d2,'k2=',k2,'k3=',k3,'k7=',k7)
```

d2= 12.639013830979696 k2= 0.00044712912811670333 k3= 0.00040451784906690475 k7= 9.7 34611256881719

C:\Users\Leandra\anaconda3\Lib\site-packages\scipy\optimize_minpack_py.py:1010: Opt
imizeWarning: Covariance of the parameters could not be estimated
 warnings.warn('Covariance of the parameters could not be estimated',

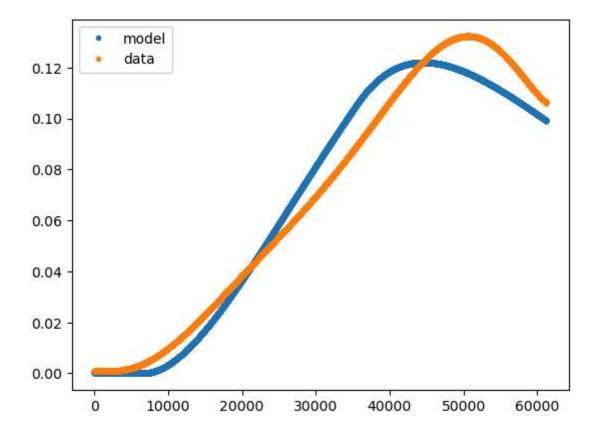
```
In [8]: import sys
        import numpy
        \#params = [1,1,1,1,1,1,1]
        params = [1,1,1,1]
        numpy.set printoptions(threshold=10)
        \#model1 = np.asarray(func(t,Kd,n,d2,k2,k3))
        model1 = np.asarray(func(t,d2,k2,k3,k7))
        print(len(model1))
        #print(model1)
        \#a,b,c,d,e,f,g = params
        a,b,c,d= params
        ydata = np.asarray(new)
        print(len(ydata))
        ssr = np.sum((ydata-model1)**2)
        #ssr2 = np.sum((ydata-model2)**2)
        sst = np.sum((ydata-np.mean(ydata))**2)
        R2=1-ssr/sst
        print('R2 is:', R2)
       51
       51
       R2 is: 0.9587598513656904
In [9]: pp = PdfPages('multipage.pdf')
        ydata = np.asarray(new)
        \#condition = [1,2,3,4,5,6,7,8,9]
```

```
In [9]: pp = PdfPages('multipage.pdf')
    ydata = np.asarray(new)

#condition = [1,2,3,4,5,6,7,8,9]
    for i in range(1):
        #model = tester(t,Kd,n,d2,k2,k3,i)
        model = tester(t,d2,k2,k3,k7,i)

#print(model)
        #a,b,c,d,e,f,g= params
        a,b,c,d= params

        plt.plot(t,model,'.', label = 'model')
        #print(model)
        #t = np.Linspace(0,34800, num=6961)
        #raw = rawdata[0:13920:240]
        plt.plot(t,rawdata,'.',label = 'data')
        #print(rawdata[i])
        plt.legend()
        pp.savefig()
        plt.show()
```



In []:
In []:
In [10]: #####