## Practical 5 - Oscillations

# Jan Izquierdo Ramos, Carmen Samedi Gamboa October 2024

Jan Izquierdo, Carmen Samedi

### SYSTEMS AND NETWORK BIOLOGY - PRACTICAL 5

#### Oscillations

To submit your report, answer the questions below and save the *notebook* clicking on File > Download as > iPython Notebook in the menu at the top of the page. Rename the notebook file to ''practicalN\_name1\_name2.ipynb'', where N is the number of the practical, and name1 and name2 are the first surnames of the two team members (only one name if the report is sent individually). Finally, submit the resulting file through the *Aula ESCI*.

Remember to label the axes in all the plots.

IMPORTANT REMINDER: Before the final submission, remember to **reset the kernel** and re-run the whole notebook again to check that it works.

The aim of this practical is to explore the dynamics of oscillatory systems and the requirements for oscillations.

[1]: # Load all necessary Python packages here
import numpy as np
from scipy.integrate import odeint
import matplotlib.pyplot as plt

#### A. Oscillations from a negative feedback: the Goodwin oscillator

We will begin by studying the following two-dimensional system:

$$\frac{dx_1}{dt} = \frac{1}{1+x_2^n} - k_1 x_1$$

$$dx_2$$

$$\frac{dx_2}{dt} = x_1 - k_2 x_2$$

- a) Consider that  $k_1 = k_2 = 0.1$ . Is there any value of n lower than 12 for which the system shows sustained oscillations?
- b) For each of the previous cases, how does the system behave on the phase plane? (Draw the nullclines and plot the trajectory of the system on the phase plane.)

```
[7]: k1=k2=0.1
     def osc(k_list, t):
         k1=k_list[0]
         k2=k_list[1]
         dx1_dt=1/(1+x2**n) - k1*x1
         dx2_dt=x1 - k2*x2
         return [dx1_dt, dx2_dt]
     time=np.linspace(0, 1000, 500)
     for n in range(12):
         d=odeint(osc, [k1, k2], time)
         print(d)
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We can see there is no value lower than 12 for which the system shows sustained oscillations.

c) Now add a third variable  $x_3$  that evolves according to:  $\frac{dx_3}{dt} = x_2 - k_3 x_3$ , with  $k_3$ =0.1, and consider that  $x_1$  is inhibited by  $x_3$  instead of  $x_2$ .

Is there now a value of n for which the system shows sustained oscillations? (Make sure you integrate for sufficiently long time, it may help to plot the beginning and the final part of the simulation in different subplots.

## []: # Write your code here

Comment your result here

d) Add saturation to the degradation of  $x_3$ , with Michaelis constant K\_d:

$$\frac{dx_3}{dt} = x_2 - \frac{k_3 \, x_3}{K_d + x_3}$$

Consider that  $k_1 = k_2 = 0.1$ ,  $k_3 = 2$  and  $K_d = 0.01$ . What do you observe now as n increases?

## []: # Write your code here

Comment your result here

B. Oscillations from a single intermediate

A negative feedback system can oscillate even with only one intermediate, if there are enough nonlinearities in the system. To see this, consider again the initial 2D system, but assume that there is an additional pathway of  $x_2$  degradation that can saturate and whose activity is inhibited by high  $x_2$  values.  $x_2$  dynamics is now described by the following ODE:

$$\frac{dx_2}{dt} = x_1 - k_2 x_2 - \frac{k_3 x_2}{K_m + x_2 + x_2^2}$$

- a) Plot the time evolution of the system for  $k_1=k_2=0.1,\,k_3=5,\,n=2,\,K_m=0.1.$
- b) Increase  $K_m$  up to 10. What happens?
- c) Plot the nullclines of the system and superimpose the corresponding trajectories. What do you observe?

[]:	# Write your code here	
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