due date Friday, Feb 11, 2022

Biochemical switch (100 points)

(Problem 3.7.5 from Strogatz' book)

As one ingredient of biological pattern formation, Lewis and coworkers (1977) considered a model of a biochemical switch, in which a gene G is activated by a biochemical substance S to produce a pigment or other gene product. Let g(t) denote the concentration of the gene product as a function of time. The model is given by

$$\frac{dg}{dt} = k_1 s_0 - k_2 g + \frac{k_3 g^2}{k_4^2 + g^2}$$

where s_0 is the (time-independent) concentration of the substance S and the k's are positive constants. The first term describes the stimulation of the production of g by the substance S, the second is a linear degradation of g and the nonlinear term is an autocatalytic (positive) feedback.

1. Show that the system can be rewritten in dimensionless form as

$$dx/d\tau = s - rx + \frac{x^2}{(1+x^2)}$$

for suitably defined dimensionless quantities x, τ and parameters s, r.

- 2. Study the the model in the absence of stimulus (i.e. s = 0). Find all fixed points and analyze their stability. Show that some fixed points only exist if the degradation is not too strong, $r < r_c$. Find r_c .
- 3. Assume that initially there is no gene product, g(0) = 0, and suppose s is slowly increased from zero (the activating signal is turned on). What happens to g(t)? What happens if s then goes back to zero; does the gene turn off again?
- 4. Consider the model with stimulus ($s \neq 0$). Study the limiting cases of small and large s. Which terms in the differential equation do you need to keep? Find the fixed points in these limiting cases.
- 5. Now study the model numerically. This can be done either using a package like MathLab or by writing a program that integrates the differential equation. Explore the entire (r, s) parameter space and give a quantitatively accurate plot of the stability diagram.

Hint: Follow the fixed points found in 2. with increasing s.