Biol635/Math635/Biol432/Math430 Fall 2024

Assignment 5

.

Answer the following questions.

- Justify your answers.
- Explain your results.
- Provide the necessary calculations in a clear way.
- Provide the codes you used (if any).

Consider the following (slightly) modified version of the *logistic equation with a threshold*

$$
\frac{dV}{dt} = F(V) \tag{1}
$$

where t represents time, V is the dependent variable (e.g., voltage) and the function $F(V)$ is given by

$$
F(V) = -rV\left(1 - \frac{V}{T}\right)\left(1 - \frac{V}{K}\right) + I.
$$
\n(2)

The parameters r, K, T and I are constants, r, K, $T > 0$ and $T < K$. The parameter r represents the rate constant (inverse of the time constant for the unbiased case when $I = 0$). The units depend on the problem at hand. If the model is intended for the dynamics of a neuron, then the units of t are milliseconds (ms) and the units of V are millivolts (mV).

This equation has many applications. In particular, it can be used to describe the transition of a system from a resting state (V_{rest} , lower equilibrium V_{eq}) to and activated state (V_{act} , higher equilibrium V_{ea}), when they exist, through a threshold (V_{th} , intermediate equilibrium V_{eq}). V_{act} is also referred to as the saturation value V_{sat} .

In the context of neuronal systems, eq. (1) can be used as a toy / phenomenological (very simple) model of action potential generation (in the absence of termination) where the occurrence of an action potential is determined by the transition from V_{rest} to V_{act} . In this case, the parameter I is interpreted as the current (I_{app}) the neuron receives (e.g., an external DC current that is controlled by the experimenter/modeler). The dynamics governing the termination of an action potential are not described by this model, and requires additional mechanisms and additional model complexity (e.g., a system of two or more differential equations).

The existence and values of the equilibria (V_{rest} , V_{th} and V_{sat}) depend on the model parameters (r, T, K and I). For $I = 0$, eq (1) has three equilibria (V_{eq}) : $V_{rest} = 0$, $V_{th} = T$ and $V_{sat} = K$. Because of this, T and K are referred as the threshold and carrying capacity, respectively, in the literature. For other values of I, the activity attributes (V_{rest} , V_{th} and V_{sat} are not longer explicitly given by a single parameter (T, K) and the attributes may even cease to exist for certain ranges of values of I (all other parameters fixed).

- 1. Write a code to solve numerically the ODE (1)-(2) (or adapt the template code provided in the course website). The simulation output for each set of parameter values must be
	- (a) A graph of the solution $V(t)$.
	- (b) The equilibrium value(s) $V_{eq} = \lim_{t\to\infty} V(t)$
	- (c) A graph of F as a function of V .
	- Each simulation should be run long enough (large enough value of t) so that $V(t)$ reaches values close enough to V_{eq} , but not too long so the changes in $V(t)$ are clearly shown and can be appreciated.
	- Plot the two graphs as two panels in the same Figure.
	- Use correct labels for the axes.
	- Use large enough fonts (suggested: "fontsize" $= 24$)
- 2. Consider the following parameter values: $r = 1, T = 0.25, K = 1$. Perform simulations as described above for $V(0) = 0.01$ and three values of $I: I = 0, I = 0.05, I = 0.1$.
- 3. Consider the following parameter values and initial condition: $r = 1, T = 0.4, K = 1, I = [0, 0.2]$ with intervals $\Delta I = 0.02$ (11 values), and $X(0) = 0.25$
	- (a) Simulate the model as described above in **ascending** order of the values of I_{app} . Plot the graph of V_{eq} as a function of I.
	- (b) Simulate the model as described above in **descending** order of the values of I_{app} . Plot the graph of V_{eq} as a function of I_{app} .
- 4. Consider the following parameter values and initial condition: $r = 1, T = 0.4, K = 1, I_{amp}$ $[0, 0.2]$ with intervals $\Delta I = 0.02$ (11 values)
	- (a) Simulate the model as described above in **ascending** order of the values of I. For $I = 0$ use $V(0) = 0$. For $I_{app} > 0$ set $V(0)$ equal to V_{eq} in the simulation for the previous value of I_{app} .
	- (b) Simulate the model as described above in **descending** order of the values of I_{app} . For $I_{app} = 0.2$ set $V(0)$ equal to the value of V_{eq} computed in the previous simulation for $I_{app} =$ 0.2. For $I_{app} < 0.2$ set $V(0)$ equal to V_{eq} in the previous simulation.
	- (c) Plot a single graph with all the values of V_{eq} as a function of I_{app} .
- 5. (Graduate level) Provide examples from the literature on bistability in neuronal systems