

Omdugga 2012

Omdugga 1, questions 1-3. Omdugga 2, questions 3-5. Omdugga both, questions 1-5.

Good luck! ☺

1 Consider the following little system:

$$\begin{aligned}d/dt(x_1) &= -k_1 \cdot x_1 + k_2 \cdot x_2 \\d/dt(x_2) &= k_1 \cdot x_1 - k_2 \cdot x_2 - k_3 \\y_{\text{hat}} &= x_1 + x_2 + k_{\text{meas}} \\x_1(0) &= 0.5, x_2(0) = 0.6 \\k_1 &= 1, k_2 = 2, k_3 = 3 \\k_{\text{meas}} &= 5\end{aligned}$$

- Which are the states in this system?
- Which are the reaction rates?
- Are any of the reactions reversible/irreversible? Why/why not?

2 Cost functions and optimization

- What is the input and output of a cost function. What does it do?
- What are the residuals? Both give a formula, and say in words what they "do".
- What is the input and output of an optimization algorithm? What does it do?

3 Consider the following system:

$$\begin{aligned}d/dt(x_1) &= -k_1 \cdot x_1 + u \\d/dt(x_2) &= k_1 \cdot x_1 - k_2 \\y_{\text{hat}} &= (x_1 + x_2) \cdot k_{\text{meas}} \\x_1(0) &= 0.5, x_2(0) = 0.6 \\k_1 &= 1, k_2 = 2 \\k_{\text{meas}} &= 5\end{aligned}$$

- Assume that the k_1 -reaction is saturated, with a Michaelis-Menten expression. What changes in the model?
- What is the residual at time $t=0$, if the measurement is $y(0) = 4$?
- What are the reactions in the following model?

$$\begin{aligned}d/dt([A]) &= k_1 - V_{\text{max}} \cdot [A] / (K_m + [A]) + k_2 \cdot [B] \\d/dt([B]) &= + V_{\text{max}} \cdot [A] / (K_m + [A]) - k_2 \cdot [B] - k_3 \cdot [B] \\y_{\text{hat}} &= k_y \cdot [A]\end{aligned}$$

4 Statistical tests:

- a) What do you conclude if you do not reject a whiteness test?
- b) What is the null hypothesis of a chi-square test?
- c) Assume that you have two acceptable models, but where one of them has a slightly lower cost than the other. How can you test whether this difference is significant? What is the test, and what should happen (reject/not reject)?

5 Closing the loop, predictions and experimental tests.

- a) What is the problem with parameters in biological models describing complex systems? How does this affect the quality of the predictions, compared to e.g. the situation in physics?
- b) Name one type of conclusion that you can draw using a model. How can that conclusion be stronger because of the model, compared to if you didn't have it, and just looked at the data?
- c) You are sitting at the table at the end of the project, discussing with the customer. Your analysis of the model(s) have shown that a certain experiment would be a good idea; e.g. measuring the amount of a protein after 10 minutes. What could be a possible reason for doing that measurement? Also specify what the corresponding model prediction would be to that argument; is it a core prediction?