

**Read.** *Mathematical Concepts in Modern Biology*, Chapter 1: Mechanisms of gene regulation: Boolean network models of the lactose operon in *Escherichia coli*, by R. Robeva, B. Kirkwood, and R. Davis, pages 1–35.

**Do.** Familiarize yourself with either Macaulay2, Singular, or Sage.

### Exercises.

- Given a Boolean network  $(f_1, f_2, f_3, f_4, f_5)$ , suppose that  $\{x_1 + x_5, x_3 + 1, x_2 + x_5 + 1, x_4\}$  is a Gröbner basis of the ideal  $I = \langle f_i + x_i \mid i = 1, \dots, 5 \rangle$ . What can you deduce from this? Your answer should consist of several clear and complete sentences, and *you should be as specific as possible*.
- Consider the following system of polynomial equations:

$$\begin{aligned}x^2 + y^2 + xyz &= 1 \\x^2 + y + z^2 &= 0 \\x - z &= 0\end{aligned}$$

- Use a software package to compute a Gröbner basis for this system over  $\mathbb{Q}$ . Include a print-out of the code that you used.
  - Use the Gröbner basis you just computed to write a simpler system of polynomial equations that has the same set of solutions. Solve that system *by hand* (it's not hard) to find all (complex-valued) solutions to the original system.
  - Repeat the previous two parts, but over the binary field,  $\mathbb{F}_2 = \{0, 1\}$ .
  - Now, do Parts (a) and (b) but over the ternary field,  $\mathbb{F}_3 = \{0, 1, 2\}$ .
- Consider the following simple model of the *lac* operon:

$$\begin{array}{ll}f_M = \overline{R} & f_R = \overline{A} \\f_P = M & f_A = L \wedge B \\f_B = M & f_L = P\end{array}$$

For this problem, make the convention that  $(x_1, x_2, x_3, x_4, x_5, x_6) = (M, P, B, R, A, L)$ .

- Sketch the wiring diagram of this Boolean network.
- Justify each function in a single sentence. What other assumptions are made in this model? (E.g., presence or absence of external lactose and glucose?)
- Write each function as a polynomial over  $\mathbb{F}_2 = \{0, 1\}$ . Then, write out a system of equations whose solutions are the fixed points of the Boolean network.
- Find the fixed points by computing a Gröbner bases with a software package.
- Compute and print out the entire phase space of your model with the help of the Cyclone algorithm on AlgoRun, at <http://algorun.org/browse>.

4. Consider the following Boolean network:

$$(f_1, f_2, f_3) = (\overline{x_1} \wedge x_2 \wedge \overline{x_3}, (x_1 \wedge \overline{x_3}) \vee (\overline{x_1} \wedge x_3), \overline{x_1} \wedge x_2 \wedge \overline{x_3}).$$

- (a) Sketch the wiring diagram of this Boolean network.
- (b) Write these functions as polynomials in  $\mathbb{F}_2[x_1, x_2, x_3]/\langle x_1^2 - x_1, x_2^2 - x_2, x_3^2 - x_3 \rangle$ . Feel free to use Macaulay2, Sage, or Singular.
- (c) Sketch the (synchronous) phase space of this Boolean network. Feel free to use Algorun.
- (d) Sketch the asynchronous phase space of this Boolean network, and find all strongly connected components.
- (e) Classify the nodes of the asynchronous phase space as transient points, cyclic attractors, or complex attractors.

5.