

Read: 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: Create an account on the Sage Math Cloud (<https://cloud.sagemath.org>).

1. Consider the following system of polynomial equations:

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

To compute a Gröbner basis for this system over \mathbb{R} , type the following commands into Sage, one-by-one, and press Shift+Enter after each one:

```
P.<x,y,z> = PolynomialRing(RR, 3, order='lex'); P
I = ideal(x^2+y^2+x*y*z-1, x^2+y+z^2, x-z); I
B = I.groebner_basis(); B
```

- (a) For the system above, 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 *real-valued* solutions to the original system.
 - (b) Are there any *complex-valued* solutions not in \mathbb{R} ? [*Hint:* Replace `RR` with `CC`].
 - (c) Now, solve the original system but over the binary field, $\mathbb{F}_2 = \{0, 1\}$. [*Hint:* Replace `RR` with `GF(2)`].
2. Repeat the previous problem for this system of polynomial equations:

$$\begin{aligned}x^2y - z^3 &= 0 \\2xy - 4z &= 1 \\z - y^2 &= 0 \\x^3 - 4yz &= 0\end{aligned}$$

3. 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 $(x_1, x_2, x_3, x_4, x_5, x_6) = (M, P, B, R, A, L)$.

- (a) 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?)

- (b) Write each function as a polynomial over $\mathbb{F}_2 = \{0, 1\}$. Then, write out the system of equations $\{f_i + x_i = 0, i = 1, \dots, 6\}$, whose solutions are the fixed points of the Boolean network.
- (c) Go into Sage and type the following command:

```
P.<x1,x2,x3,x4,x5,x6> = PolynomialRing(GF(2), 6, order='lex'); P
```

Now, define an ideal I generated by the six polynomials from Part (b). Use Sage to compute the Gröbner basis of this ideal, and include a print-out of a screenshot.

- (d) The Gröbner basis describes a simpler system of equations with the same solutions as the original. Write out this system and then solve it by hand to determine the fixed points of the Boolean network.
- (e) Compute the entire phase space of your model with the help of the Analysis of Dynamic Algebraic Models (ADAM) toolbox, at <http://adam.plantsimlab.org/>. Under “Model Type”, select Polynomial Dynamical System (PDS). Print a screenshot of the state space graph. Are there any periodic points that are not fixed points?