

## Final Exam Review Problems

For details on exam coverage and a list of topics, please see the class website. Note that this is a set of review problems, and NOT a practice exam. Most problems below concern material covered after Exam 2. Please see the review problems for Exams 1 and 2 for a more detailed review of material from earlier in the course.

1. Consider a chemical reaction network with master equation

$$\frac{dp_k}{dt} = \sum_{j=1}^m \rho_j^\sigma(k - \gamma_j) p_{k-\gamma_j} - \sum_{j=1}^m \rho_j^\sigma(k) p_k$$

Suppose that the initial distribution satisfies

$$\sum_{k \in \mathbb{Z}_+^n} p_k(0) = 1$$

(That is, the initial distribution is really a distribution). Show that for any time  $t \geq 0$ ,

$$\sum_{k \in \mathbb{Z}_+^n} p_k(t) = 1$$

That is, that the solutions of the CME are always a probability distribution, assuming that the initial conditions were.

2. Assume that mRNA is produced in “bursts” of  $r > 1$  transcripts at a time. Further, as before, mRNA degrades.
  - (a) Write down a system of chemical reactions describing the above phenomena.
  - (b) Find the stoichiometry matrix of the network.
  - (c) Assuming mass-action kinetics, write down the propensity functions.
  - (d) Write the CME for the bursting model.
3. A population of 10 individuals is divided into two types,  $A$  and  $a$ . Let  $X(t)$  denote the number of type  $A$ s at step  $t$ . There is a separate reservoir of individual in which the frequency of  $A$  is always 0.4. The population evolves as follows: with probability 0.1 an individual is removed at random from the population of 10 and replaced by a randomly drawn individual from the reservoir; with probability .9 the population is updated as in the Moran model, that is, two individuals are drawn at random with replacement and the second is replaced by a copy of the first.

Find

$$p_{i,i-1} = \mathbb{P}(X(t+1) = i-1 \mid X(t) = i),$$

for  $i = 1, 2, \dots, 10$ .

4. State what the infinite population assumption says about the frequency of genotypes among children of a mating population.
5. Let  $A$  be the state transition matrix for a Markov chain evolving on the state space  $\{0, 1, 2, 3, 4\}$ .

$$A = \begin{pmatrix} 1/6 & 5/6 & 0 & 0 & 0 \\ 1/6 & 1/2 & 1/3 & 0 & 0 \\ 0 & 2/3 & 1/6 & 1/6 & 0 \\ 0 & 0 & 1/3 & 1/3 & 1/3 \\ 0 & 0 & 0 & 1/3 & 2/3 \end{pmatrix}$$

You can also compute the following powers of  $A$ :

$$A^2 = \begin{pmatrix} 1/6 & 5/9 & 5/18 & 0 & 0 \\ 1/9 & 11/18 & 2/9 & 1/18 & 0 \\ 1/9 & 4/9 & 11/36 & 1/12 & 1/18 \\ 0 & 2/9 & 1/6 & 5/18 & 1/3 \\ 0 & 0 & 1/9 & 1/3 & 5/9 \end{pmatrix}, \quad A^3 = \begin{pmatrix} 0.12 & 0.60 & 0.23 & 0.05 & 0 \\ 0.12 & 0.55 & 0.26 & 0.05 & 0.02 \\ 0.09 & 0.52 & 0.23 & 0.10 & 0.06 \\ 0.04 & 0.22 & 0.20 & 0.23 & 0.31 \\ 0 & 0.07 & 0.13 & 0.32 & 0.48 \end{pmatrix}$$

$$A^4 = \begin{pmatrix} 0.12 & 0.56 & 0.25 & 0.05 & 0.02 \\ 0.11 & 0.55 & 0.24 & 0.07 & 0.03 \\ 0.10 & 0.49 & 0.24 & 0.09 & 0.08 \\ 0.04 & 0.27 & 0.18 & 0.22 & 0.29 \\ 0.01 & 0.12 & 0.15 & 0.29 & 0.42 \end{pmatrix}$$

At time  $t = 0$ ,

$$(\mathbb{P}(X(0) = 0), \mathbb{P}(X(0) = 1), \mathbb{P}(X(0) = 2), \mathbb{P}(X(0) = 3), \mathbb{P}(X(0) = 4)) = \left(0, \frac{2}{3}, \frac{1}{6}, 0, \frac{1}{6}\right).$$

- (a) Draw a state transition diagram for this chain.
  - (b) Is the chain irreducible? Why or why not?
  - (c) Find  $\mathbb{P}(X(8) = 3 \mid X(4) = 1)$ .
  - (d) Compute  $\mathbb{P}(X(3) = 4)$ .
  - (e) Find  $\mathbb{E}(X^2(4) \mid X(3) = 3)$ .
  - (f) Find  $\lim_{t \rightarrow \infty} \mathbb{P}(X(t) = 3)$ .
6. Suppose  $X$  takes on each of the values 1, 2 and 3 with probability  $1/3$ .
- (a) Find the moment-generating function of  $X$ .
  - (b) Find the cumulant-generating function of  $X$ .
  - (c) Using your result from part (b), compute the mean and variance of  $X$ .
7. Consider a population of individuals able to produce offspring of the same kind, so that each individual, by the end of its lifetime, will have produced  $j$  new offspring with probability  $p_j$  for  $j = 0, 1, 2, \dots$ , independently of the numbers produced by the other individuals. Clearly then,

$$0 \leq p_j \leq 1,$$

$$\sum_{j=0}^{\infty} p_j = 1.$$

Suppose that there exists **one individual in the initial generation** (generation 0), and let  $X(t)$  denote the size of the  $t^{\text{th}}$  generation, for  $t = 0, 1, 2, \dots$ . More formally,

$$X(t) = \sum_{i=1}^{X(t-1)} Z_i$$

where  $Z_i$  denotes the number of offspring of the  $i^{\text{th}}$  individual of the  $(t-1)^{\text{st}}$  generation, and the  $Z_i$  have distribution described by the previous  $p_j$ .

Let

$$\mu := \mathbb{E}[Z_i] = \sum_{j=0}^{\infty} j p_j$$

In HW #8, we showed that if  $\mu < 1$ , then

$$\lim_{t \rightarrow \infty} \mathbb{P}(X(t) = 0 \mid X(0) = 1) = 1,$$

i.e. that the population dies out with probability 1. Assume now that  $\mu > 1$ .

- (a) Recall that  $\pi_0 = \mathbb{P}(\text{population eventually dies out})$ , since 0 is an absorbing state. Assume  $X(1) = j$ . Find the following probability:

$$\mathbb{P}(\text{population eventually dies out} \mid X(1) = j)$$

- (b) By conditioning on  $X(1)$ , find an equation satisfied by  $\pi_0$ . *Note:* It can be shown that  $\pi_0$  is the smallest positive number satisfying the correct equation for this part.
- (c) Suppose that  $p_0 = 1/2, p_1 = 1/4, p_2 = 1/4$ . Find  $\pi_0$ .
- (d) Suppose that  $p_0 = 1/4, p_1 = 1/4, p_2 = 1/2$ . Find  $\pi_0$ .
8. In class, when discussing the Luria-Delbrück distribution, we assumed that the resistant cells grew deterministically:

$$Y_n^N = \frac{N}{n} X_n,$$

or

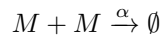
$$Y_n^N = \left(\frac{N}{n}\right)^c X_n,$$

in the case of relative growth rate  $c$ . Now assume that the resistant mutants grow stochastically, so that

$$Y_n^N = K_n^N X_n,$$

where  $K_n^N$  is the **random** size of a mutant population at phage application first appearing when there are  $n$  sensitive cells. That is,  $K_n^N$  is a random variable. All other dynamics are assumed identical to the case discussed in class.

- (a) Find an expression for the cumulant generating function of  $Z$ , the number of mutants when the phage is applied. *Note:* This will depend on the statistical properties of the random variable  $K_n^N$ .
- (b) Find the mean and variance of  $Z$ .
9. Consider the annihilation of a species  $A$  via dimerization:



- (a) Write down the chemical master equation for this reaction.
- (b) What do you expect the stationary distribution  $\pi$  of this reaction network to be?
- (c) Prove your claim from part (b).