Problem Set 3: Networks and Gene Regulation

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1 Bayesian Decision Theory

2 Gibbs Sampling For Motif Discovery

My Gibbs sampling algorithm is modeled after the algorithm described in Chapter 12.2 of An Introduction to Bioinformatcs Algorithms. It also makes use of relative entropy to weigh the random choice of a starting position (step 5 as described on page 413). I make use of pseudocounts for my profile matrix. I do **not** use the log-of-odds when calculating my l-mer probabilities, as Clojure handles extremely small decimal numbers fairly well.

My metric for convergence is a value referred to in the code as the *cutoff*. For simplicity, I say the algorithm has converged when cutoff rounds go by without an increase in the maximum-probability l-mer. I've arbitrarily set the cutoff to be 20 rounds in the current implementation, but it can easily be adjusted. If I had more time, it would be fun to explore using simulated annealing instead of my naive approach.

Despite using relative entropy, my solution seems to perform poorly when the AT/GC ratio is non-uniform. This suggests to me that I am not making proper use of the relative-entropy equation.

Regardless, below is a table summarizing the most frequent 10-mer motifs discovered across the four datasets provided:

Data 1	Data 2	Data 3	Data 4
ATTCGAATTC	GTCTACTACT	AAAAAAAAA	AAAAAAAAA
TCGAATTCCC	CTACTACTCA	TTTTTTTTT	AACAAAAAA
TTCGAATTCC	TCATATAACA	AAAAAAAAGA	TTTTTTTTT
CGAATTCGAA	CTGTCTACTA	AAAAAAAACA	AAAAAAAAAT
AATTCGAATT	TCTCTTAAGA	TTGTATATAT	ATAAATAAAT

As you can see, the results for data 3 and 4 are likely not actual motifs.