Practice final for Math 5233. You are allowed to use your notes for the exam, but not the textbook or printed articles. This exam is longer than the actual exam in order to illustrate a variety of possible questions.

- (1) What is the difference between a transversion and a transition between nucleotides? What is the significance of this for bioinformatics?
- (2) Using a scoring system of +3 for a match, -2 for a mismatch, and -1 for a gap, use the Smith-Waterman algorithm to fill in a dynamic programming table and find the best local alignment between the sequences AACTG and ACCG. If there are ties, list all of the best alignments.
- (3) Calculate the score (in bits) of matching the sequence ACGG with a PSSM derived from the aligned sequences
 - ACGG ACAG ACAA ТСАА

assuming a uniform background distribution of nucleotides.

- (4) What are G-protein coupled receptors? What techniques can be used to recognize them from their amino acid sequences?
- (5) The PROSITE pattern for G6PD (glucose-6-phosphate dehydrogenase) is D - H - [YF] - L - G - K - [EQK].How many amino acid sequences of length 7 match this pattern? How many amino acid sequences of length 7 are there? If you found a match to this pattern in the Macaca mulatta (rhesus monkey) genome, how confident would you be that it was orthologous to G6PD? What could you do to check?
- (6) Using the complete mitochondrial genomes, the Jukes-Cantor distances in the table below were calculated. Use the UPGMA algorithm to construct a possible phylogeny for these 5 species.

	Chimp.	Gorilla	Orang.	Human	Pigmy Chimp
Chimp.	0	80	120	65	30
Gorilla	80	0	120	80	80
Orangutan	120	120	0	120	120
Human	65	80	120	0	65
Pygmy Chimp.	30	80	120	65	0



FIGURE 1.

- (7) What are the assumptions of the Jukes-Cantor model? Do you think that sites in the multiple DNA sequence alignment in Figure 1 satisfy those assumptions? Why or why not?
- (8) Explain why the statement "protein A and protein B are 50 percent homologous" is incorrect.
- (9) Suppose the entries of a DNA positional scoring matrix are computed from the formula $s(a, b) = \log_2(16f_{ab})$, where f_{ab} are empirical frequencies of DNA substitutions (a is the column, b is the nucleotide). How would you interpret an entry equal to -5? What assumptions are being made about the statistical properties of the sequence?
- (10) True or False: it does not matter which program you use to compute a maximum parsimony phylogenetic tree you will get the same answer as long as you use the same data, since there is a unique optimal solution. Explain your answer.
- (11) Compute the Burrows-Wheeler transform of the string 'AARDVARK'.
- (12) Is the alignment below from a protein-coding region of DNA, an RNA with a conserved secondary structure, or the mitochondrial D-loop? Explain your reasoning.



- (13) What are the implicit assumptions of the Nussinov folding algorithm (the version used in our worksheet)? How could you modify the algorithm to be more realistic?
- (14) The VGF protein is known to be extremely important in the brain, being involved in many different processes. However, it is not very well understood. In particular, it has no known conserved domains. If you wanted to find a conserved structure in this protein, what mathematical tools could you use?
- (15) Describe an algorithm that would efficiently find the lowest cost path from the top to the bottom of a pyramid of numbers such as the one shown below. The cost is the sum of the numbers along the path. All paths must start from the top and go through a number immediately below and to the left or the right. An example path is shown with a cost of 47. Use pseudo-code (i.e. a precise outline) to describe exactly what the algorithm would do.

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