ID: 105667404

CSCI 3104 Problem Set 7 Profs. Grochow & Layer Spring 2019, CU-Boulder

1. Multiple string alignment. As in class, we consider the operations of substitution (including the zero-cost substitute-a-letter-for-itself "no-op"), insertion, and deletion. An alignment of three strings x, y, z (of lengths n_x, n_y, n_z , respectively) is an array of the form:

That is, in the first row x appears in order, possibly with some gaps, in the second row y does, and in the third row z does. We define the cost of such an alignment as follows. The cost of a column

 y_1

is the sum-of-pairs cost, e.g., in the preceding example we look at the cost of aligning x_2 with y_1 (a substitution, costing either 0 or 1 depending on whether $x_2 = y_1$ or not), the cost of aligning x_2 with — (a deletion, costing 1), and the cost of aligning y_1 with — (another deletion), for a total cost of $c_{subs} + 2c_{del} = 3$ for this one column of the alignment. The total cost of the alignment is the sum of the costs of its columns. Given three strings x, y, z, the goal of Multiple String Alignment is to find a minimum-cost alignment.

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(a) (15 pts) Give an efficient (polynomial-time) algorithm for finding optimal alignments of three strings. Describe your algorithm in pseudocode and English, and prove an upper bound on its running time; you do not need to prove correctness (but you will only receive full credit if your algorithm is correct!). Hint 1: start with a recursive algorithm, where the recursion has 7 cases depending on what the last column of the alignment looks like:

$$\begin{pmatrix} x_{n_x} \\ y_{n_y} \\ z_{n_z} \end{pmatrix}, \begin{pmatrix} - \\ y_{n_y} \\ z_{n_z} \end{pmatrix}, \begin{pmatrix} x_{n_x} \\ - \\ z_{n_z} \end{pmatrix}, \begin{pmatrix} x_{n_x} \\ y_{n_y} \\ - \end{pmatrix}, \begin{pmatrix} - \\ - \\ z_{n_z} \end{pmatrix}, \begin{pmatrix} - \\ y_{n_y} \\ - \end{pmatrix}, \begin{pmatrix} x_{n_x} \\ - \\ - \end{pmatrix}$$

Hint 2: Your recursive algorithm will likely not be very efficient; what technique can you use from class to improve its efficiency?

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(b) (7 pts) Consider generalizing your approach to part (a) to k strings instead of 3. How many cases would there be to consider in the recursion? What runtime would the algorithm have? You do not need to give the algorithm, but must argue persuasively that your answers are correct, given your answer to part (a).

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2. The most efficient version of the preceding approach we know of for three strings takes an amount of time which becomes impractical as soon as the strings have somewhere between 10⁴ and 10⁵ characters (which are still actually fairly small sizes if one is considering aligning, e. .g, genomes). Because of this, people seek faster heuristic algorithms, and this is even more true for aligning more than three strings. One natural approach to faster multi-string alignment is to first consider optimal pairwise alignments and somehow use this information to help find the multi-way alignment. Here we show that naive versions of such a strategy are doomed to fail.

Given an alignment of three strings, we may consider the 2-string alignments it induces: just consider two of the rows of the alignment at a time. If both of those rows have a gap in some column, we treat the pairwise alignment as though that column doesn't exist. For example, the alignment of x and y induced by the figure at the top of Q1 would be

Note that the column between x_3 and y_3 got deleted because it consisted of two gaps.

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(a) (5pts) Give an example of three strings x, y, z such that the optimal alignment of x with y does not appear in any optimal alignment of x, y, z, and the same is true for optimal alignments of x, y and of y, z. Prove your example has this property. (An example where any optimal alignment of x, y, z does not contain an optimal alignment of x, y—but may contain optimal alignments of x, z and/or y, z—will earn you partial credit.)

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(b) (7 pts) Prove that the gap between the cost of the pairwise optimal alignments and the pairwise alignments induced by an optimal 3-way alignment can be arbitrarily large. More symbolically, let d_{xy} denote the cost of an optimal alignment of x and y, and for an alignment α of x, y, z, let $d_{xy}(\alpha)$ denote the cost of the pairwise alignment of x and y which is induced by α . Your goal here is: for any c > 0, construct three strings x, y, z such that $d_{xy}(\alpha) > d_{xy} + c$ for any three-way alignment α . (Suppose all three strings have length n; how big can you make the gap c as a function of n, asymptotically?)

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3. (14 pts) Give an efficient algorithm to compute the number of optimal solutions to the Knapsack problem. Recall the Knapsack problem has as its input a list $L = [(v_1, w_1), \ldots, (v_n, w_n)]$ and a threshold weight W, and the goal is to select a subset S of L maximizing $\sum_{i \in S} v_i$, subject to the constraint that $\sum_{i \in S} w_i \leq W$. For this problem, you will count the number of such optimal solutions. Your algorithm should run in O(nW) time. If you only give an inefficient recursive algorithm, you can still receive up to 6 pts. (If you are having trouble solving this problem, you may want to start by developing an inefficient recursive algorithm and then seeing how to improve it using techniques from class.)

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4. Consider the following variant of string alignment: given two strings x, y, and a positive integer L, find all contiguous substrings of length at least L that are aligned (using no-ops = substituting a letter for itself) in some optimal alignment of x and y. Assume the costs of substitution, insertion, and deletion are given by constants $c_{subs}, c_{ins}, c_{del}$, and that the cost of substituting a letter for itself is zero.

(a) (2 pts) Show that the output here contains at most $O((n-L)^3)$ substrings.

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(b) (10 pts) Give an algorithm that solves this problem. You may use/modify/refer to the pseudocode for Knapsack from the lecture notes.