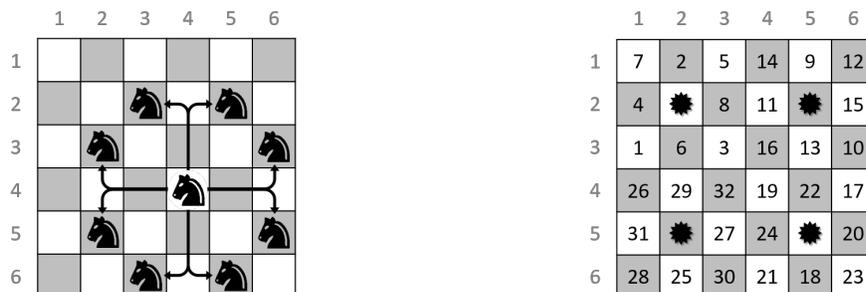


# Homework #1

( Due: Oct 6 )



## Task 2. [ 50 Points ] Distance-Incorporated Codon Autocorrelation Score

The availability of synonymous codons (codons that can translate the same amino acid into a protein) enables a protein to be encoded by many different sequences of codons/tRNAs. Autocorrelation measures the reuse of a particular codon/tRNA in succession (instead of choosing a different synonymous one) during the translation of a protein sequence. Studies show that tRNA autocorrelation in a coding sequence has important effects on its translation speed.

We can measure autocorrelation by transforming the problem into a combinatorial one. For example, suppose we have a sequence of amino acids where Serene's residues occur at positions 4, 6, and 301, and to be coded by a mix of codons from two different tRNAs (say, two of type A and one of type B). These codons can appear in three different relative orders: AAB, BAA, and ABA. Considering the opportunities of tRNA reuse, AAB is expected to translate faster and is more autocorrelated than BAA, because the two occurrences of A in AAB are so close that the specific tRNA molecule employed in coding for the 4th residue is likely to be around when residue at position 6 is being translated, and hence can be reused.

*Distance Incorporated Codon Autocorrelation* (DICA) score is a recently proposed metric of gene autocorrelation which is calculated for a coding sequence by finding positions of all synonymous codons for a given amino acid and then summing a reward function,  $F(d(i, j))$  which assigns a positive score based on the distance between the synonymous codons. Here,  $d(i, j)$  is the distance between codons translated by the same tRNA, i.e., if a tRNA repeat is found at positions  $i$  and  $j$  then the distance between these two is  $d(i, j) = j - i$ . Because autocorrelation appears to decay slowly with distance, for a given synonymous codon, the probability that the next codon is the same decreases as the distance increases. It has been shown that an exponential distance function of the form  $c^{d(i, j)}$  where  $c \approx 0.999$  gives a better measure for DICA.

Suppose we have a codon sequence  $\mathcal{S}$  of length  $n$  composed of  $K$  ( $\leq 9$ ) different amino acids<sup>2</sup>  $\mathcal{A}_a$ ,  $1 \leq a \leq K$ . Let  $\mathcal{S}_a$  be the maximal subsequence of  $\mathcal{S}$  containing only  $\mathcal{A}_a$ 's, and let  $n_a$  be its length. Then  $n = \sum_{a=1}^K n_a$ , and clearly,  $n = \Theta(\max_{a=1}^K n_a)$ . For any given  $a \in [1, K]$  and  $1 \leq i \leq n_a$ , let  $l_i^{(a)}$  be the location of  $\mathcal{S}$  where the  $i$ -th entry of  $\mathcal{S}_a$  occurs. Then the *Distance-Incorporated Codon Autocorrelation* (DICA) score ( $D_{\mathcal{S}}$ ) for  $\mathcal{S}$  is:

$$D_{\mathcal{S}} = \frac{\sum_{a=1}^K \sum_{i=1}^{n_a} \sum_{j=i+1}^{n_a} \theta(T_{l_i^{(a)}}, T_{l_j^{(a)}}) \times F(d(l_i^{(a)}, l_j^{(a)}))}{\sum_{a=1}^K \sum_{i=1}^{n_a} \sum_{j=i+1}^{n_a} F(d(l_i^{(a)}, l_j^{(a)}))},$$

where  $T_i$  is the tRNA at location  $i$  of  $\mathcal{S}$ ,  $\theta(T_i, T_j) = 1$  if  $T_i = T_j$  and 0 otherwise. Note that there can be at most  $m$  different tRNA's, where  $m \leq 6$ .

This equation can be evaluated using three nested loops. However, such a naïve computation of DICA takes  $\Theta(n^2)$  time, which can be prohibitively expensive for very long sequence.

Show how to compute  $D_{\mathcal{S}}$  for a sequence of length  $n$  in  $\Theta(n \log n)$  time. Include pseudocode of your algorithm.

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<sup>2</sup>There are 9 amino acids with synonymous codons translated by different tRNAs ( $A, G, I, L, P, R, S, T, V$ )

SELECT(  $A[q : r]$ ,  $k$ ,  $d$ ,  $s_{\text{even}}$ ,  $s_{\text{odd}}$ ,  $b$  )

**Input:** An array of distinct elements, and an integer  $k \in [1, r - q + 1]$ . The parameter  $d$  is the depth of recursion with  $s_{\text{even}}$  being the block size to be used at even depths and  $s_{\text{odd}}$  at odd depths. Also  $b$  is an upper bound on the size of the base case.

**Output:** An element  $x$  of  $A[q : r]$  such that  $\text{rank}(x, A[q, r]) = k$ .

1.  $n \leftarrow r - q + 1$
2. **if**  $n \leq b$  **then**
3.     sort  $A[q : r]$
4.     **return**  $A[q + k - 1]$
5. **else**
6.     **if**  $d \bmod 2 = 0$  **then**  $s \leftarrow s_{\text{even}}$
7.     **else**  $s \leftarrow s_{\text{odd}}$
8.     divide  $A[q : r]$  into blocks  $B_i$ 's each containing  $s$  consecutive elements  
       ( last block may contain fewer than  $s$  elements )
9.     **for**  $i \leftarrow 1$  **to**  $\lceil \frac{n}{s} \rceil$  **do**
10.          $M[i] \leftarrow$  median of  $B_i$  using sorting
11.      $x \leftarrow$  SELECT  $\left( M[1 : \lceil \frac{n}{s} \rceil], \left\lfloor \frac{\lceil \frac{n}{s} \rceil + 1}{2} \right\rfloor, d + 1, s_{\text{even}}, s_{\text{odd}}, b \right)$      {median of medians}
12.      $t \leftarrow$  PARTITION(  $A[q : r]$ ,  $x$  )     {partition around  $x$  which ends up at  $A[t]$ }
13.     **if**  $k = t - q + 1$  **then return**  $A[t]$
14.     **else if**  $k < t - q + 1$  **then return** SELECT(  $A[q : t - 1]$ ,  $k$ ,  $d + 1$ ,  $s_{\text{even}}$ ,  $s_{\text{odd}}$ ,  $b$  )
15.     **else return** SELECT(  $A[t + 1 : r]$ ,  $k$ ,  $d + 1$ ,  $s_{\text{even}}$ ,  $s_{\text{odd}}$ ,  $b$  )

Figure 2: Selection with hybrid blocking.

### Task 3. [ 60 Points ] Recursive Selection with Hybrid Blocking

Figure 2 shows a slightly generalized version of the selection algorithm we saw in the class. Instead of using a single block size (e.g., 5) at all levels of recursion, it uses block size  $s_{\text{even}}$  at even levels, and  $s_{\text{odd}}$  at odd levels. Now the base case size  $b$  is also a parameter to the algorithm. Observe that when  $b = 140$  and  $s_{\text{even}} = s_{\text{odd}} = 5$ , the algorithm reduces to the one we saw in the class.

- (a) [ 10 Points ] Write a recurrence relation describing the running time of SELECT on an array of size  $n$  assuming  $s_{\text{even}} = s_{\text{odd}} = 3$ . What is the best running time you get by solving the recurrence? What is the smallest value of  $b$  you get?
- (b) [ 20 Points ] Repeat part (a) with  $s_{\text{even}} = 3$  and  $s_{\text{odd}} = 5$ .
- (c) [ 30 Points ] Suppose we run SELECT with  $s_{\text{even}} = s_{\text{odd}} = 4$ . Then in steps 8–10, each group of size 4 will have exactly 2 medians — one smaller and one larger. Suppose in step 10, we assign the smaller median to  $M[i]$ . Then what will be the running time of SELECT on an array of size  $n$ ? If the running time is  $\omega(n)$ , can one make change(s) to steps 11–15 to bring the running time down to  $\Theta(n)$ ?