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CSE4074 Computer Networks Project#2

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INTRODUCTION

We worked on a particular region in a genome, so we only took some part of the genome as input. This input defined as one line of text in a file where the text contained only A, T, G or C. In project document, input file has 10 lines, where each line contains strings of length 500.

Our objective is to search for motifs and try to find the consensus string by using Randomized Motif Search and Gibbs Sampler.

After we found the consensus strings, we compared the scores and consensus strings obtained for different k values and gave it as output.

RANDOMIZED ALGORITHMS

Randomized algorithms make random rather than deterministic decisions. The main advantage is that no input can reliably produce worst-case results because the algorithm runs differently each time. These algorithms are commonly used in situations where no exact and fast algorithm is known. There are 5 different most used randomized algorithms:

- Randomized Quick-Sort
- Randomized Algorithms
- Greedy Profile Motif Search
- Gibbs Sampler
- Random Projections

RANDOMIZED MOTIF SEARCH

Randomized Motif Search may change all t motifs in $Motifs = (Motif_1, ..., Motif_t)$ in a single iteration. This strategy may prove reckless, since some correct motifs may potentially be discarded at the next iteration.

```
Algorithm Randomized Motif Search

RandomizedMotifSearch(DNA, k, t)

1: randomly select k-mers Motifs = (Motif_1, ..., Motif_t), one from each string in DNA

2: BestMotifs \leftarrow Motifs

3: while forever do

4: Profile \leftarrow Profile(Motifs)

5: for i \leftarrow 1 to t do

6: Motif_i \leftarrow Profile-most probable k-mer in the i-th string in DNA

7: Motif_i \leftarrow Motif_i \cdots Motif_t

8: if Score(Motifs) < Score(BestMotifs) then

9: BestMotif_s \leftarrow Motifs

10: else

11: return BestMotif_s
```

(Figure-1: Pseudocode of Randomized Motif Search)

GIBBS SAMPLER

Gibbs Sampler is a more cautious iterative algorithm that discards a single k-mer from the current set of motifs at each iteration and decides to either keep it or replace it with a new one. It starts with randomly chosen k-mers in each of t DNA sequences, but it makes a random rather than a deterministic choice at each iteration. It uses randomly selected k-mers (Motif₁, ..., Motif_t) to come up with another (hopefully higher scoring) set of k-mers. It randomly selects an integer i between 1 and t and then randomly changes only one Motif_i.

Algorithm Gibbs Sampler

```
GibbsSampler(DNA, k, t, N)

1: randomly select k-mers Motifs = (Motif_1, \dots, Motif_t) in each string in Dna

2: BestMotifs \leftarrow Motifs

3: \mathbf{for} \ j \leftarrow 1 \ \text{to} \ N \ \mathbf{do}

4: i \leftarrow \text{Random}(\mathbf{t})

5: Profile \leftarrow \text{profile} \ \text{matrix} \ \text{constructed} \ \text{from all strings} \ \text{in} \ Motifs \ \text{except} \ \text{for} \ Motif_i \leftarrow \text{profile-randomly} \ \text{generated} \ k-mer in the i-th sequence in DNA

7: if Score(Motifs) < Score(BestMotifs) \ \mathbf{then}

8: BestMotifs \leftarrow Motifs

9: return BestMotifs
```

(Figure-2: Pseudocode of Gibbs Sampler)

CODE AND OUTPUTS

(Figure-3: Sample Input File with the total number of characters)

```
kmer = seq[start:start+k] #take string with k size
def probKmerInProfile( kmer, profile ): #probability of kmer
s_curr = Score(curr_motifs, k) #find score of motifs
```

```
return dna[index][point: k + point]
def randomMotifs(dna, k):
def randomKmer( seq. k ):
def countsFromMotifs( motifs, k, initCount ): #find counts of motifs
   return mutated
def mutation(line):
   l=list(range(10)) # to get random mutation points in ten-mer we used 0 to 9 numbers list
random.shuffle(1) # we shuffled the list to get first four of the list as random positions different than each other
 ef Score ( motifs, k ):
```

```
def findConsensus(bestMotifs,k):
            if (profile[i][j]>maxProb): #take the base with max probability from the profile
       consensus.append(indexToBase(base)) with convert base index to base, and add chosen base to the consensus
print("Please enter the file path: " ) #take dna file name as an input
print("please enter the k value") #take k-mer size value as input
bestMotifs1 = randomizedMotifSearch(k, dna) feall randomized motif search for dna array from the input file and k value
```

(Figure-4: Codes of the Project)

```
Please enter the file path:
C///Bears//Eeysa//Desktop//Input.txt

please enter the k value

Results of Randomized Motif Search

###Best Motifs##

GGTGGCGCT

GGGGGCTCT

CGGGGCTCT

CGGTGCACA

TGTGGCTCC

CGGGCTCT

CGGGGCTCT

CGGGGCTCA

CGGGCTCA

CGGGCTCA

Consensus String: CGGGGCTCA

Score: 23

Results of Gibbs Sampler

###Best Motifs##

AACTCATCG

AACTCTCAG

AACACAAC

TACTCAGAC

AACACAAC

TACTCACAG

AACCCCCCG

GGCCCACAG

ATCTCACAG

ATCTCACAG

CACCCACAG

ATCTCACAG

CACCCACAG

ATCTCACAG

CACCCACAG

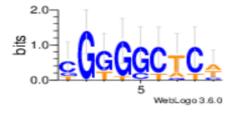
ATCTCACAG

ACCTCACAG

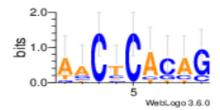
COnsensus String: AACTCACAG

Score: 21
```

(Figure-5: Output of k is 9)



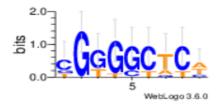
(Figure-6: Consensus string of Randomized Motif Search for k is 9)



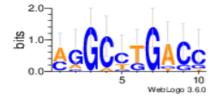
(Figure-7: Consensus string of Gibbs Sampler for k is 9)

```
Please enter the file path:
please enter the k value
Results of Randomized Motif Search
###Best Motifs###
CGGGTAGATT
TCGATTGCTC
Score:
###Best Motifs###
AGGCCTGACT
AGGAGTGAGC
AAGCCTGTCC
AGGCCTGACG
```

(Figure-8: Output of k is 10)



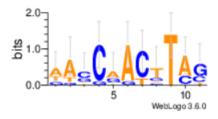
(Figure-9: Consensus string of Randomized Motif Search for k is 10)



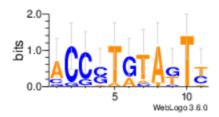
(Figure-10: Consensus string of Gibbs Sampler for k is 10)

```
Please enter the file path:
please enter the k value
Results of Randomized Motif Search
###Best Motifs###
AGGCAACATCC
TTCCGACGTAC
AAACCACTTAG
TATCAGCTTAG
AACCTATTTAG
###Best Motifs###
GGCCTATATTT
ACCGTGTATTC
ACGGTGTTGTC
CCCGTGTATTA
ACCGTGAAGTC
Consensus String:
```

(Figure-11: Output of k is 11)



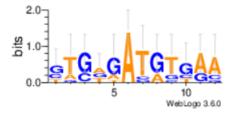
(Figure-12: Consensus string of Randomized Motif Search for k is 11)



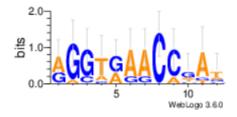
(Figure-13: Consensus string of Gibbs Sampler for k is 11)

```
Please enter the file path:
###Best Motifs###
TTCAGATGTGAC
CTCAGATAGTGG
CTGATATATGAA
GTGAGATGCCGA
GTGGGACATCAA
GTGTAATGGGAA
GTGGGAGGTGAA
                    GTGAGATGTGAA
Results of Gibbs Sampler
###Best Motifs###
GGGTGAACCGAA
GGCTGGACCGAG
AGGTGAACCAGT
AGGTGAACCGAG
                     AGGTGAACCGAT
```

(Figure-14: Output of k is 12)



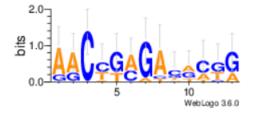
(Figure-15: Consensus string of Randomized Motif Search for k is 12)



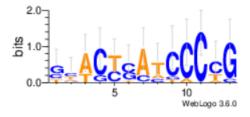
(Figure-16: Consensus string of Gibbs Sampler for k is 12)

```
Please enter the file path:
Results of Randomized Motif Search
###Best Motifs###
AACTTCGACATGA
AACCGAGTCAATG
AGCTTAGACTAGG
###Best Motifs###
TATCCGACCCCCG
CTACCCATCCCCG
CCTGTTATCCCCG
                     GCACTCATCCCCG
```

(Figure-17: Output of k is 13)



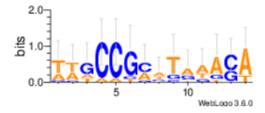
(Figure-18: Consensus string of Randomized Motif Search for k is 13)



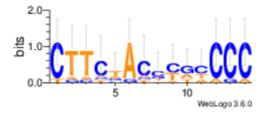
(Figure-19: Consensus string of Gibbs Sampler for k is 13)

```
Results of Randomized Motif Search
###Best Motifs###
GCGCCGGCTAAGGA
TTGCCTCTTAGACA
AAAACGCATAAACA
TATCAGCTTAGACT
ATTCCCCAGGAACA
Consensus String:
Results of Gibbs Sampler
###Best Motifs###
CTTGTATCCGACCC
CTCAGACCCACCCC
CTTTTACGTGCACC
CTTCTACCCATCCC
CATCAAGCTTCCCC
CTTCAACGTGCCCC
CTTCGGCTTGACCC
CTTCTACTATTCGA
Consensus String:
                     CTTCTACCCGCCCC
```

(Figure-20: Output of k is 14)



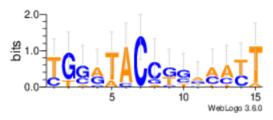
(Figure-21: Consensus string of Randomized Motif Search for k is 14)



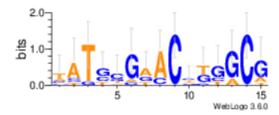
(Figure-22: Consensus string of Gibbs Sampler for k is 14)

```
Please enter the file path:
please enter the k value
Results of Randomized Motif Search
###Best Motifs###
CTCGTCCCGTGAATT
TGGGTACCGGACATT
TGCAAACCTCAACTT
TGGCTACCTGTTACT
                   TGGATACCGGAAATT
Results of Gibbs Sampler
###Best Motifs###
AATGGGTACCGGACA
TATGTAAACTTTGCG
AATGTGGACGTGGCG
CGTGAGACCATTGCG
Consensus String:
                     TATGCGAACCTGGCG
```

(Figure-23: Output of k is 15)



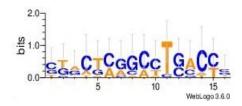
(Figure-24: Consensus string of Randomized Motif Search for k is 15)



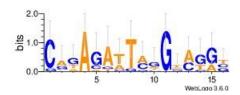
(Figure-25: Consensus string of Gibbs Sampler for k is 15)

```
Please enter the file path:
Results of Randomized Motif Search
###Best Motifs###
CGTCAAGGCCTGACTT
CTACTCAGACCCACCC
GGGCGAGCCCTCATCG
TTGCGCGGCTTGACTA
GCGCGCGGCCTGGCCG
CTAATCGGCTTCAACG
CGACTACCCATCCCTC
GTCTTCGGCTTGACCC
Results of Gibbs Sampler
###Best Motifs###
CGGACATTAGGTCGAG
CAAAGATTCGGAATGC
CGGAAGTTAAGTAGGT
CATAGTTTGGGCATGT
CCGAGAATCTGTCGGG
CTTAGACTAGGGCGGG
GGTAGATTATGTAGGT
CAAAGATTCGGCACAT
CAGACAATCCGGAGGC
                     CAGAGATTAGGTAGGT
                      54
```

(Figure-26: Output of k is 16)



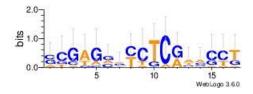
(Figure-27: Consensus string of Randomized Motif Search for k is 16)



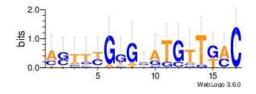
(Figure-28: Consensus string of Gibbs Sampler for k is 16)

```
Please enter the file path:
please enter the k value
Results of Randomized Motif Search
###Best Motifs###
GGCTGGATCTCGTCCCG
GCGTAAACCTCGATGCT
ACGAGGGTCTCACACCG
CCGACAACTGCAAACCT
GTCAAGTTCTCGACCTT
GCGTGAGTTTCGGCCCT
Consensus String:
Results of Gibbs Sampler
###Best Motifs###
TCTTCGGTTATGTTTCC
AGAGCGGGGCTCTTGAC
ACTTTGCGCGGCTTGAC
AGTTTGGGCATGTTTCC
ACCATGTGAATGTGGAC
AGGGCGGGGGTGTTGAC
CGTCTAAGTATGTTGAC
CGTTCGGGAATGATTCC
                     AGTTTGGGCATGTTGAC
```

(Figure-29: Output of k is 17)



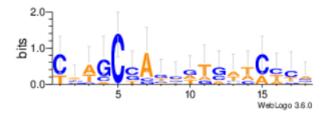
(Figure-30: Consensus string of Randomized Motif Search for k is 17)



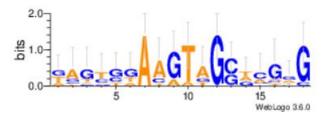
(Figure-31: Consensus string of Gibbs Sampler for k is 17)

```
Please enter the file path:
Results of Randomized Motif Search
###Best Motifs###
CATCCCAGCTGGTTCTTG
CGAGCCCTCATCGTCTCT
TCAGCTATCTTCTACCCA
CTTGCGACAGACCTCCTA
CATGCTACGTTGACACAC
CTCCCCATGATGATATTC
TCAGCCAGTATAACCCCA
Consensus String:
                     CAAGCCAGCGTGATCCCA
Results of Gibbs Sampler
###Best Motifs###
GAAGTTAAGTAGGTCGTG
TAGCCGAAGTTGCACGAG
ATGTGGACGTGGCGCGCG
GATTGGAAGTAGGAATAG
TAGATTATGTAGGTAGAG
```

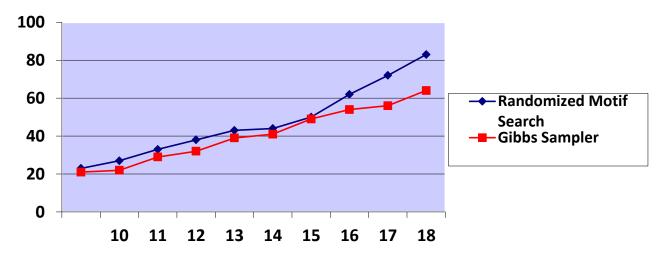
(Figure-32: Output of k is 18)



(Figure-33: Consensus string of Randomized Motif Search for k is 18)



(Figure-34: Consensus string of Gibbs Sampler for k is 18)



(Graph-1: Score of Randomized Motif Search and Gibbs Sampler for Different 'k' Values)

CONCLUSION

In this project, we used a txt file (input.txt) that has 10 lines, where each line contains strings of length 500 that include randomly generated genomic bases. Then, we selected a random position between 0-490 to insert the 10-mer that has 4 mutations in 4 random positions. In our programs, we take a k value which is the length of the consensus string.

In Randomized Motif Search, we chose a random motif with length k for each line. Then, we calculated the score with initial count zero for each base and the profile matrix of them. By choosing best motif, we found new motifs from DNA strings and calculated score and the profile matrix of new motifs. We repeated these processes until it believes that there is no more best score out there. It keeps best score value when one score comes higher than the best score algorithm halts and returns its best value so, it caught up by local optima. Finally, we had 10 motifs and found the consensus string using these.

In Gibbs Sampler, we chose a random motif with length k for each line and selected a random x point to take xth motif off from motifs. Then, we calculated the score with initial count one for each base and the profile matrix of them after deleting xth motif. By choosing a random motif from the xth DNA with using probabilities, we found new motifs from DNA strings and calculated score and the profile matrix of new motifs. We repeated these processes until the score not change in the last 50 iterations. Finally, we had 10 motifs and found the consensus string using these.

After, we implemented Randomize Motif Search and Gibbs Sampler, we get information that Gibbs Sampler randomly tries all possibilities and gives chance to all bases by taking initial count of bases, so it is better to see every chance and gives better score than Random Motif Search. However, Randomized Motif Search works faster than Gibbs Sampler because it returns best motif that it finds the first. Lastly, we noticed that the scores of Randomize Motif Search and Gibbs Sampler are mostly increasing as the k-mers' size increased.

At the end of the program, we have 10 motifs for each algorithm. According to these motifs' count probability, we found the consensus string and checked them by using

"http://weblogo.threeplusone.com/create.cgi" link. Then, we added all outputs and consensus strings in our report.

REFERENCES

http://bix.ucsd.edu/bioalgorithms/presentations/Ch12 RandAlgs.pdf

 $\underline{https://www.cs.helsinki.fi/u/tpkarkka/teach/16-17/MAiB/MAiB2016-lecture2.pdf}$

http://www.mit.edu/~ilkery/papers/GibbsSampling.pdf