## CS 640: Hidden Markov Models

## Generalized Hidden Markov Models are defined as follows:

- 1. N hidden states: S1,,...,SN
- 2. M symbols in emitted alphabet
- 3. Initial probability distribution vector of length N:  $\mathbf{\pi} = \{ \pi_1 ..., \pi_N \}$
- 4. Transition probability matrix of size N x N

where  $\tau_{ij}$  is probability of transition from state in row i to state in row j

5. Emission probability matrix of size N x M where **e**<sub>i</sub>(c) probability that state i emits character c

We refer to the transition probabilities, the emission probabilities and the initial distribution vector, collectively as the parameters of the model, designated  $\lambda = (\tau_{ij}, e_i(c), \pi)$ .

Let Q be the sequence of visited states:  $Q = (q_1, q_2, ..., q_F)$ Let O be the sequence of emitted symbols:  $O = (O_1, O_2, ..., O_T)$  (the observed sequence).

Write a generalized Hidden Markov Model that employs the Viterbi algorithm (which is a dynamic programming algorithm) to find most likely sequence of hidden states to emit an observed sequence. You may hard-code in a transition matrix, emissions matrix and start probabilities. Your program should read a string of any length in FASTA format from a file and output the score of the nucleotide sequence of that string, given the HMM defined below. The string may contain whitespace and numbers, which should be ignored, as well as nucleotide characters {A, C, T, G}

Code the Viterbi algorithm for this HMM, filling in matrix cells  $\alpha_t(\mathbf{i})$ , where t corresponds to sequence index and i corresponds to state:

- 1. N = 3. hidden states S<sub>1</sub>, S<sub>2</sub>, S<sub>3</sub>
- 2. M=4 symbols in alphabet {a, c, t, g}
- 3. Initial probability distribution vector  $\pi = \{.25, .5, .25\}$
- 4. Transition probability matrix **τ** =

	<b>S</b> 1	S2	<b>S</b> 3
<b>S</b> 1	.5	.4	.1
S2	0	.5	.5
<b>S</b> 3	.3	.2	.5

## 5. Emission probabilities **e** =

	a	c	t	g
<b>S</b> 1	.4	.3	.2	.1
S2	.25	.25	.25	.25
<b>S</b> 3	.1	.2	.3	.4

Initialization:  $\alpha_1(i) = \pi_i e_i(O_1)$ 

Iteration:  $\alpha_{t+1}(i) = e_i (O_{t+1})_j \max_{j \in \text{ states}} (\alpha_t(j) * \tau_{ji})$ 

Sean Eddy generalized HMMS:http://www.nature.com/nbt/journal/v22/n10/full/nbt1004-1315.html