# CS 466 Introduction to Bioinformatics Lecture 20 

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## Course Announcements

Project Proposal due Nov 14th

## Hidden Markov Model $\mathcal{M}=(Q, A, \Sigma, E)$

- Set of hidden states $Q$
- Markov property
- Transition probabilities $A=\left[a_{i j}\right]$ on pairs of states
- Set of emitted symbols $\Sigma$
- Emission probabilities $E=\left[e_{i k}\right]$ on state-symbol pairs

Two decisions:

1. What symbol should I emit? [emission probabilities $E$ ]
2. What state should I move to next? [transition probabilities $A$ ]

Fair Bet Casino

$$
\begin{array}{rl}
Q & =\{F, B\} \\
F & B \\
A & =\left(\begin{array}{cc}
0.9 & 0.1 \\
0.1 & 0.9
\end{array}\right)_{B}^{F} \\
\Sigma & =\{H, T\} \\
H & T \\
E & =\left(\begin{array}{cc}
0.5 & 0.5 \\
0.75 & 0.25
\end{array}\right)_{B}^{F}
\end{array}
$$



## Three Questions

## Question 1:

What is the most probable path $\boldsymbol{\pi}^{*}$ that generated observations $\mathbf{x}$ ?

## Question 2:

What is probability of observations $\mathbf{x}$ generated by any path $\boldsymbol{\pi}$ ?

## Question 3:

What is the probability of observation $x_{i}$ generated by state $s$ ?

| Type | Probability | Method | Solution |
| :--- | :--- | :--- | :--- |
| Joint | $\max _{\boldsymbol{\pi} \in Q^{n}} \operatorname{Pr}(\mathbf{x}, \boldsymbol{\pi})$ | Viterbi algorithm | $\operatorname{Pr}\left(\mathbf{x}, \boldsymbol{\pi}^{*}\right)=\max _{s \in Q} v[s, n]$ |
| Marginal | $\operatorname{Pr}(\mathbf{x})$ | Forward algorithm, or back- <br> ward algorithm | $\operatorname{Pr}(\mathbf{x})=\sum_{s \in Q} f[s, n]$, <br> $\operatorname{Pr}(\mathbf{x})=\sum_{s \in Q} a_{0, s} \cdot e\left[s, x_{1}\right] \cdot b[s, 1]$ |
| Posterior | $\operatorname{Pr}\left(\pi_{i}=s \mid \mathbf{x}\right)$ | Forward algorithm, and <br> backward algorithm | $\hat{\pi}_{i}=\underset{s \in Q}{\arg \max } \frac{f[s, i] \cdot b[s, i]}{\sum_{t \in Q} f[t, n]}$ |

Table 1: Hidden Markov Models - Three different probabilities

| Type | Recurrence |
| :--- | :--- |
| Viterbi | $v[s, i]=\left\{\begin{array}{ll\|}a_{0, s} e_{s, x_{1}}, \\ e_{s, x_{i}} \max _{t \in Q} v[t, i-1] \cdot a_{t, s}, & \text { if } i>1 .\end{array}\right.$ |
| Forward | $f[s, i]= \begin{cases}a_{0, s} e_{s, x_{1}}, & \text { if } i=1, \\ e_{s, x_{i}} \sum_{t \in Q} f[t, i-1] \cdot a_{t, s}, & \text { if } i>1 .\end{cases}$ |
| Backward | $b[s, i]= \begin{cases}1, & \text { if } i=n, \\ \sum_{t \in Q} a_{s, t} \cdot e_{t, x_{i+1}} \cdot b[t, i+1], & \text { if } 1 \leq i<n,\end{cases}$ |

Table 2: Hidden Markov Models - Three recurrences that each can be computed in $O\left(n|Q|^{2}\right)$ time and $O(n|Q|)$ space.


| Viterbi | $1(\mathrm{~T})$ | $\mathbf{2 ( H )}$ | $3(\mathrm{~T})$ | $\mathbf{4}(\mathrm{H})$ | $5(\mathrm{H})$ | $6(\mathrm{H})$ | $7(\mathrm{~T})$ | $8(\mathrm{H})$ | $9(\mathrm{H})$ | $10(\mathrm{H})$ | $11(\mathrm{~T})$ |
| :--- | :--- | :--- | :--- | :--- | :--- | :--- | :--- | :--- | :--- | :--- | :--- |
| F |  |  |  |  |  |  |  |  |  |  |  |
| B |  |  |  |  |  |  |  |  |  |  |  |


| Forward | 1 (T) | 2 (H) | 3 (T) | 4 (H) | 5 (H) | 6 (H) | 7 (T) | 8 (H) | 9 (H) | 10 (H) | 11 (T) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| $F$ |  |  |  |  |  |  |  |  |  |  |  |
| B |  |  |  |  |  |  |  |  |  |  |  |


| Backward | 1 (T) | 2 (H) | 3 (T) | 4 (H) | 5 (H) | 6 (H) | 7 (T) | 8 (H) | 9 (H) | 10 (H) | 11 (T) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| F |  |  |  |  |  |  |  |  |  |  |  |
| B |  |  |  |  |  |  |  |  |  |  |  |

## Questions:

1. Compute most likely state path
2. Compute marginal probability
3. Compute posterior decoding
$Q=\{F, B\}$
$A=\left(\begin{array}{ll}0.9 & 0.1 \\ 0.1 & 0.9\end{array}\right)_{B}^{F}$
$\Sigma=\{H, T\}$
$E=\left(\begin{array}{cc}{ }^{H} & T \\ 0.5 & 0.5 \\ 0.75 & 0.25\end{array}\right)_{\text {B }}^{F}$


| Type | Recurrence |
| :--- | :--- |
| Viterbi | $v[s, i]= \begin{cases}a_{0, s} e_{s, x_{1}}, \\ e_{s, x_{i}} \max _{t \in Q} v[t, i-1] \cdot a_{t, s}, & \text { if } i>1 .\end{cases}$ |
| Forward | $f[s, i]= \begin{cases}a_{0, s} e_{s, x_{1}}, & \text { if } i=1, \\ e_{s, x_{i}} \sum_{t \in Q} f[t, i-1] \cdot a_{t, s}, & \text { if } i>1 .\end{cases}$ |
| Backward | $b[s, i]= \begin{cases}1, & \text { if } i=n, \\ \sum_{t \in Q} a_{s, t} \cdot e_{t, x_{i+1}} \cdot b[t, i+1], & \text { if } 1 \leq i<n,\end{cases}$ |

## Summary

- Markov property - Current state depends only on previous state
- Hidden Markov Models: states are not given only emitted symbols
- Viterbi algorithm: Find the most likely sequence of states given a set of observations
- Baum-Welch algorithm: EM-algorithm to learn $\boldsymbol{A}$ and $\boldsymbol{E}$ from training set


## Reading:

- Jones and Pevzner: Chapters 11.1-11.3
- Lecture notes

