

Conditional  
probability

Bayes'  
Theorem

Markov  
Chain

Hidden  
Markov  
Model

# Hidden Markov Models

Garrett M. Dancik, PhD

The **conditional** probability of an event  $A$ , given that  $B$  has occurred, is denoted by  $P(A|B)$  and has the formula

$$P(A|B) = \frac{P(B|A)P(A)}{P(B)}$$

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Find the probability that a die lands on 4, given that the die roll is even.

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$$P(4|\text{even}) =$$

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$$\begin{aligned} P(4|\text{even}) &= \frac{P(\text{even}|4)P(4)}{P(\text{even})} \\ &= \frac{(1) \times \frac{1}{6}}{1/2} \end{aligned}$$



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If the event  $A$  can be broken down into disjoint events  $A_1$ ,  $A_2$ , etc, then **Bayes' Theorem** states that

$$P(A|B) = \frac{P(B|A)P(A)}{\sum P(B|A_i)P(A_i)}$$

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## Example

	M	F	Total
Soph	4	2	6
Junior	3	2	5
Senior	3	1	4
Total	10	5	15

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For some calculations, we do not need to worry about the denominator from Bayes' Theorem, and instead can work with the formula

$$P(A|B) \propto P(B|A)P(A)$$

where  $\propto$  means "is proportional to."

## Example

A sophomore student is selected. Is the student more likely to be a male or a female? Using the above version of Bayes Theorem:

$$\begin{aligned}P(M|Soph) &\propto P(Soph|M)P(M) \\&\propto \left(\frac{4}{10}\right)\left(\frac{10}{15}\right) \\&\propto \approx 0.267\end{aligned}$$

and

$$\begin{aligned}P(F|Soph) &\propto P(Soph|F)P(F) \\&\propto \left(\frac{2}{5}\right)\left(\frac{5}{15}\right) \\&\propto \approx 0.133\end{aligned}$$

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Since  $P(M|Soph) \propto 0.267$  and  $P(F|Soph) \propto 0.133$ , the selected individual is  $\frac{0.267}{0.133} = 2$  times as likely to be a male than a female.

A **Markov chain** is a sequence of random variables (or states)  $X_1, X_2, X_3$ , with the property that the next state  $X_{n+1}$  depends on the  $m$  previous states (including the current one). Usually,  $m$  is taken to be 1 in which case the next state depends only on the current one, and the Markov chain is said to have the Markov property and is a first order Markov model. Formally, first order Markov models have the property

$$P(X_{n+1}|X_1, X_2, \dots, X_n) = P(X_{n+1}|X_n)$$

## Example

An individual has two coins, a fair (F) coin and a biased (B) coin. Before each coin toss, there is a 10% chance that the individual will switch coins. Initially, there is a 50% chance the individual selects the fair (or biased) coin. Find the probability that the selected coins are FFB. Note that this is a 1st order Markov Chain. The subscript  $i$  will be used for the  $i^{th}$  selected coin.

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$$P(F_1 F_2 B_3) = P(F_1)P(F_2|F_1)P(B_3|F_2, F_1)$$



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A **Hidden Markov model** (HMM) is a Markov chain where the states are hidden (unobserved), but hidden states emit observed values with certain probabilities. The hidden states can then be deduced based on the observed values.

For a first order HMM with observed states  $O_1, O_2, \dots, O_n$  and hidden states  $H_1, H_2, \dots, H_n$ , the HMM is characterized by the following probabilities:

Initial state probabilities:	$P(H_1)$
Transition state probabilities:	$P(H_{i+1} H_i)$ for all $i$
Emission probabilities:	$P(O_i H_i)$ for all $i$

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The goal of a HMM is to predict the set of hidden states (such as the unknown gene structure). The most likely sequence of hidden states is the state sequence  $H_1, H_2, \dots, H_n$  that maximizes

$$P(H_1, \dots, H_n | O_1, \dots, O_n) \propto P(O_1 | H_1)P(H_1) \times P(O_2 | H_2)P(H_2 | H_1) \times \dots \times P(O_n | H_n)P(H_n | H_{n-1})$$

Because these probabilities are often extremely small, it is often useful to work on the log scale and find the state sequence that maximizes

$$\log P(H_1, \dots, H_n | O_1, \dots, O_n) = \log P(O_1 | H_1)P(H_1) + \log P(O_2 | H_2)P(H_2 | H_1) + \dots + \log P(O_n | H_n)P(H_n | H_{n-1}) + k,$$

where  $k$  is a constant which is not needed for our purposes.

## Coin flipping example

We will use the following probabilities:

$$P(F_1) = P(B_1) = 0.50$$

$$P(F_i|F_{i-1}) = P(B_i|B_{i-1}) = 0.90, i > 1$$

$$P(F_i|B_{i-1}) = P(B_i|F_{i-1}) = 0.10, i > 1$$

$$P(h|F) = P(\text{tail}|F) = 0.50$$

$$P(h|B) = 0.80$$

$$P(t|B) = 0.20$$

Suppose we flip 3 coins and observe  $(h, t, h)$ . Which is more likely, BFB or BBB?

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Continuing this example, suppose we knew that the two optimal states for the first two observations, ending with F or ending with B, were the following: FF and BB. There are 4 possibilities for the last hidden state:



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The optimal hidden state sequence is the above possibility that maximizes the overall probability.

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This lends itself to a dynamic programming solution (known as the **Viterbi algorithm**).

## Example

Using the HMM described previously, what sequence of coins most likely generated  $h, t, h$ ?

# High (H) vs. Low (L) GC content example

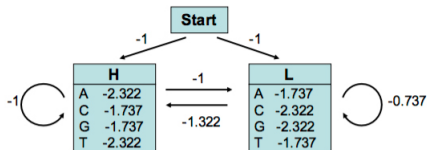
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## HMM : Viterbi algorithm - a toy example



**GGCACTGAA**

### back-tracking

(= finding the path which corresponds to the highest probability, -24.49)

	G	G	C	A	C	T	G	A	A
H	-2.73	-5.47	-8.21	-11.53	-14.01	...			-25.65
L	-3.32	-6.06	-8.79	-10.94	-14.01	...	...	...	-24.49

The most probable path is: **HHHLLLLL**

Its probability is  $2^{-24.49} = 4.25\text{E-}8$   
(remember that we used  $\log_2(p)$ )

Note: probabilities are on the  $\log_2$  scale

Source: <http://homepages.ulb.ac.be/~dgonze/TEACHING/viterbi.pdf>