

Jim Thomas  
jht@u.washington.edu  
Foege S340B

Slides and problem sets will be posted on the same web site...

Weeks 6 and 7

Intro to molecular evolution - what is it?

Neutral evolution and mutation

Purifying selection

Week 8

Phylogeny and molecular methods

Deep branches

Lineage sorting and hybridization, coalescent

Week 9

Positive (Darwinian) selection

Positive selection dN/dS methods

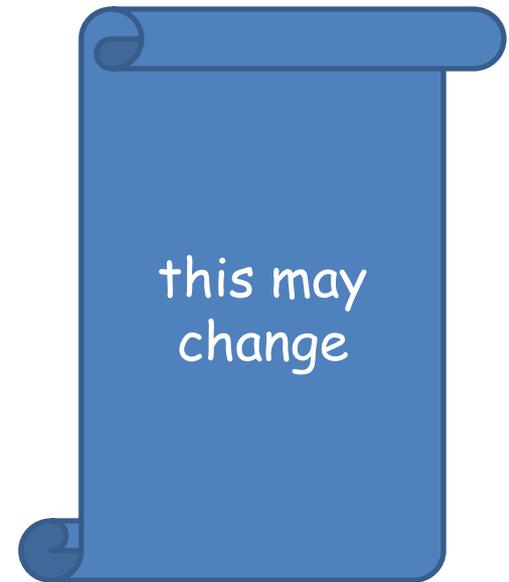
Positive selection population methods

Week 10

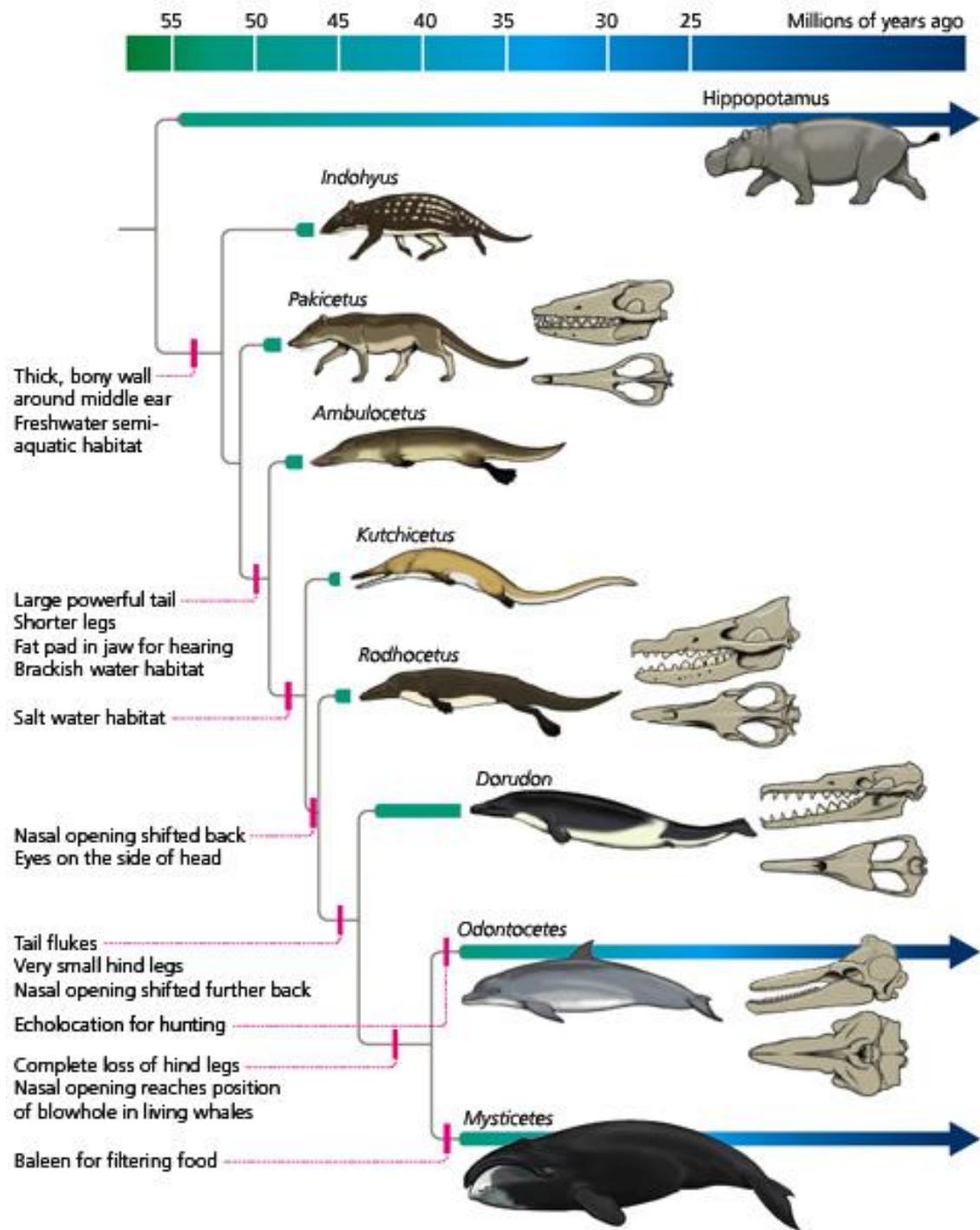
Positive selection examples 1

Positive selection examples 2

Connecting genotype and phenotype - the challenge



# Evolution of whales in the fossil record



# Molecular Evolution - What is it?



# The two fundamentals of evolution

1) heritable variation

2) natural selection

# Heritable variation

Phenotypically familiar:



hair color  
hair curliness  
eye color  
ear lobes (attached or not)  
freckling  
hair line (widow's peak or not)  
cleft chin



At least 10 genes, three of which have major effects:

EYCL1 - green/blue alleles - chromosome 19

HERC2 - central brown ring eye color gene - chromosome 15

OCA2 - brown/blue alleles - chromosome 15

PLUS, more than 2 common alleles with functional consequences for each gene

# What about molecular variation?

Molecular variation - only one kind - change in DNA sequence (or RNA for some viruses)

MRCA about 25 mYA (most recent common ancestor)



*Papio hamadryas* (baboon)



*Nomascus leucogenys* (gibbon)

```
FACATCACAATGCCAGCCAGACACTCAATAATGATGAGCTGTGCCAGTGGACACCAGAGGAGGGAAGAAAAGGAAAAGGACAAAAGTGGAGAGGGAGGAGATGGAGCAATCTTTTATGT!  
FACATCAATAACCCAGCCAGACACTCAATAATGATGAGCTGTGCCAGTGGACACCAGAGGAGGGAAGAAAAGGAGAAGGACAAAAGTGGAGAGGGAGGAGATGGAGCAATCTTTTATGT!
```



(other sequences are other primates, mostly from older branches)

```
ACATCACAATACCCT-CCCGACACTCAGTAATAATGAGCTGTGCCAGTGGACACCAGAGGAGGGGAGAAAAAGGAGGAGGACAAAAGTGGAGAGGGAGGAGATGGAGCAATCCCTTATGT!  
ACATCACAATACCCTGGCCGGACACTCAATAATGACAGCTGTGCCAGTGGACACCAGAGGAGGGGAAGAAAAAGGAGGAGGACAAAAGTGGAGCGGGAGGAGATGGAGCAGTCCCTTATGT!  
ACATCACAATACCCTGGCCAGACGCTCAATAATGACGAGCTGTGCCAGTGGACACCAGAGGAGGGGAAGAAAAAGGAGGAGGACAAAAGTGGAGCGGGAGGAGATGGAGCAGTCCCTTATGT!  
ACATCACAATACCAGCCAGACACTCAGTAGTGATGAGCTGTGCCAGTAGACACCAGAGGAGGGGAAGAAAAAGGAGAAGGACAAAAGTGGAGAGGGAAAGAGATGGAGCAATCTCTTATGT!  
ACATCACAATACCAGCCAGACACTCAATAAGTGACGAGCTGTGCCAGTAGACACCAGAGGAGGGGAAGAAAAAGGAGAAGGACAAAAGTGGAGAGGGAAAGAGATGGAGCAATCTCTTATGT!  
ACATCACAATGCCAGCCAGACACTCAATAATGATGAGCTGTGCCAGTGGACACCAGAGGAGGGGAAGAAAAAGGAAAAGGACAAAAGTGGAGAGGGAGGAGATGGAGCAATCTTTTATGT!  
ACATCACAATGCCAGCCAGACACTCAATAATGATGAGCTGTGCCAGTGGACACCAGAGGAGGGGAAGAAAAAGGAAAAGGACAAAAGTGGAGAGGGAGGAGATGGAGCAATCTTTTATGT!  
ACATCATAATACCAGCCAGACACTCAATAATGATGAGCTGTGCCAGTGGACACCAGAGGAGGGGAAGAAAAAGGAGAAGGACAAAAGTGGAGAGGGAGGAGATGGAGCAATCTCTTATGT!
```

# Natural Selection

Phenotypically familiar:

- Selection for long neck in giraffes
- Selection for bill type in Darwin's finches
- Selection for antibiotic resistance
- Selection for camouflage coloration
- Selection against sterility

etc.

What about selection on molecular variants?

What about selection on molecular variants?

Acts via the capacity for a sequence variant to affect its own perpetuation.

Recommended reading - "The Selfish Gene" by Richard Dawkins