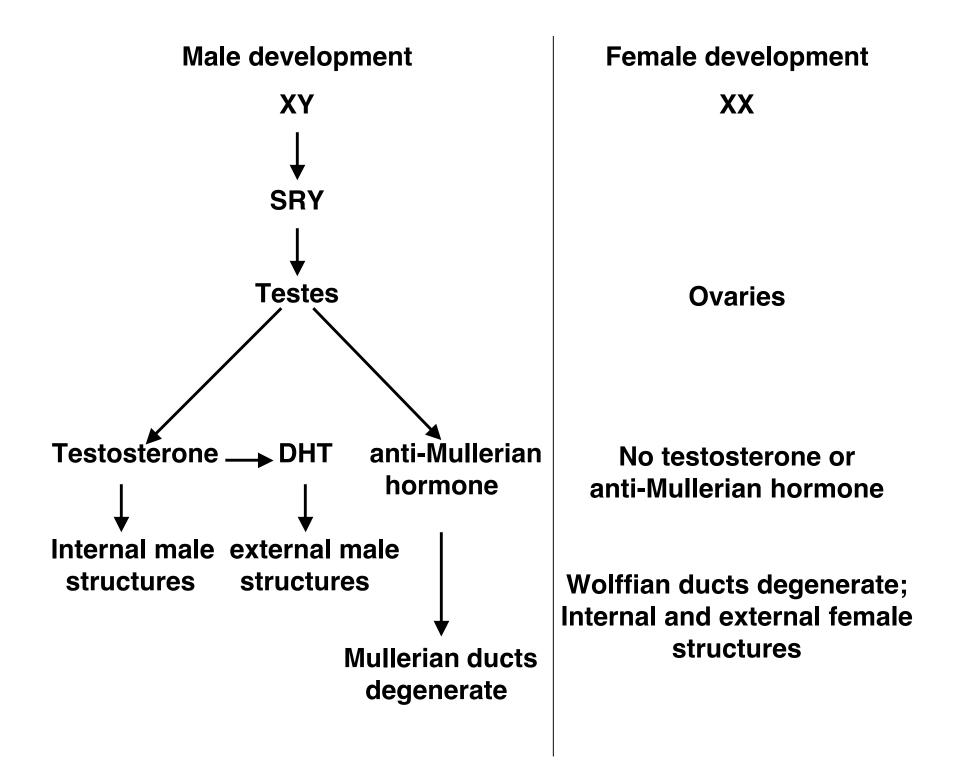
# Outline

**Review sex determination Genetic syndromes Sex-specific behaviors** sexual orientation and identity **Flies** Rodents Sheep Humans



# Mutations that affect sexual phenotypes

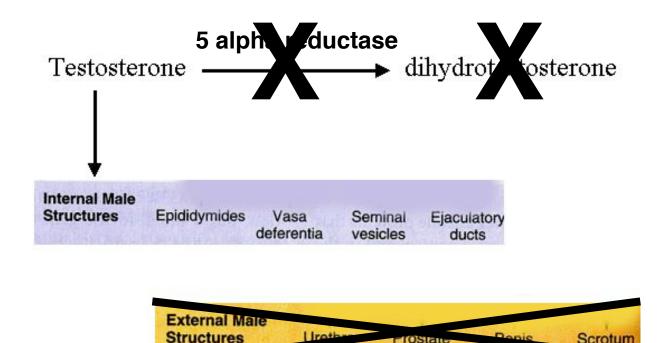
### **5-alpha-reductase deficiency**

Androgen insensitivity syndrome

**Congenital adrenal hyperplasia** 

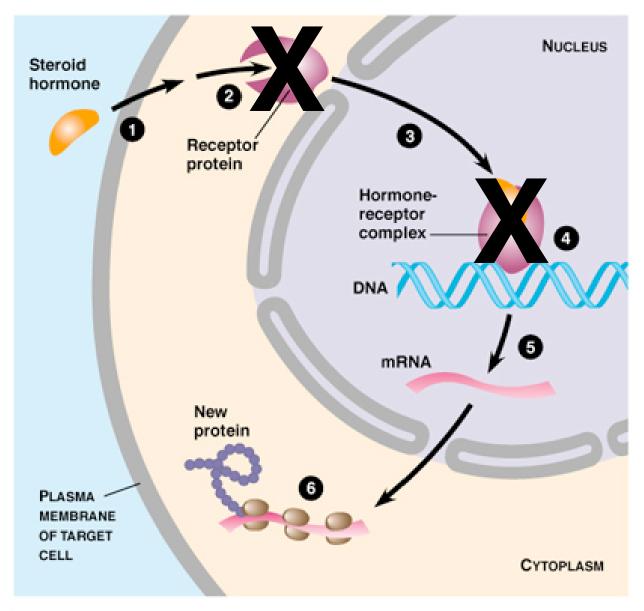
### **5-alpha-reductase deficiency**

XY individuals that lack 5 alpha reductase have testes that produce AMH and testosterone, but not DHT. AMH causes Mullerian ducts to degenerate, and testosterone induces Wolffian duct formation, but external genitalia are often female (or more female than male). At puberty massive amounts of testosterone cause the individuals sex organs to c



Scrotun

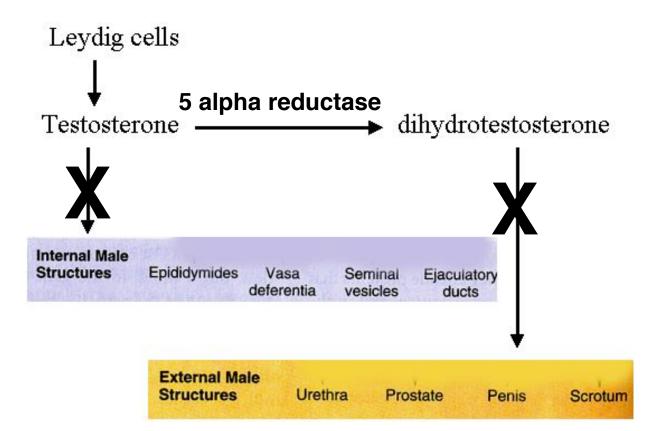
#### Androgen insensitivity syndrome (AIS)



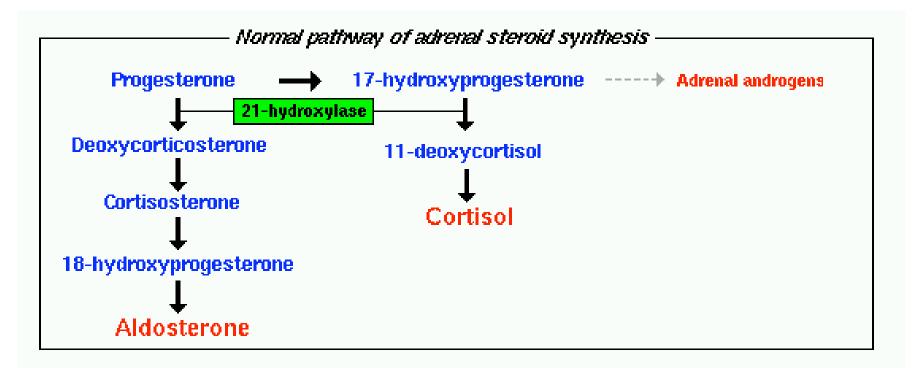
<sup>©1999</sup> Addison Wesley Longman, Inc.

### Androgen insensitivity syndrome (AIS)

XY individuals can't respond to androgens (testosterone or DHT). Testes form and produce AMH, so Mullerian ducts degenerate, but tissues don't respond to androgens, so Wolffian duct degenerates and external genitalia develop as female structures.



## **Congenital adrenal hyperplasia** Enzyme that produces aldosterone and cortisol is missing. XX individuals are masculinized.





# Outline

**Review sex determination** 

**Genetic syndromes** 

# Sex-specific behaviors

# sexual identity and orientation

## Flies

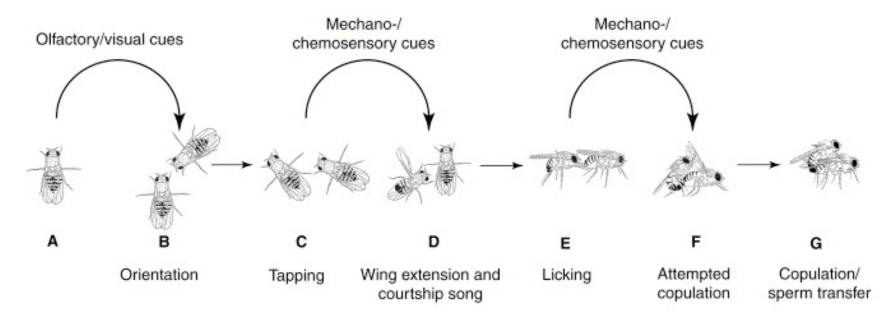
**Rodents** 

Sheep

Humans

# Sexual orientation Sexual identity

# Let's start with flies



Current Biology

### Drosophila sexual development

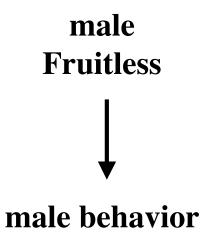
XX (X:A=1.0)

Sex lethal gene ON female Fruitless

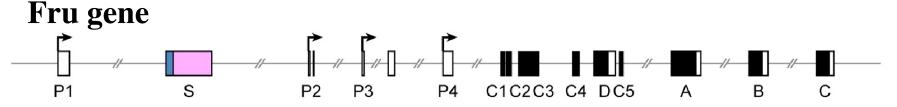
female behavior

XY (X:A=0.5)

Sex lethal gene OFF

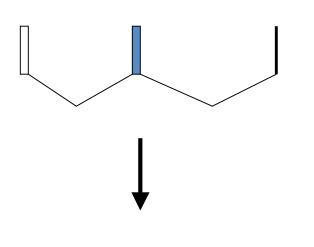


# Drosophila Fruitless (Fru) is regulated by alternate RNA splicing









female Fru protein

male Fru protein

# Fruitless controls male sexual behavior!

Fruitless mutant males lack courtship behavior.

Females that express male Fruitless exhibit male courtship behavior.

Fruitless is a transcription factor

# Females expressing male Fruitless exhibit male mating behavior.



# Male and female flies fight differently



# males box

females push

# Male and female fighting behavior is regulated by Fruitless

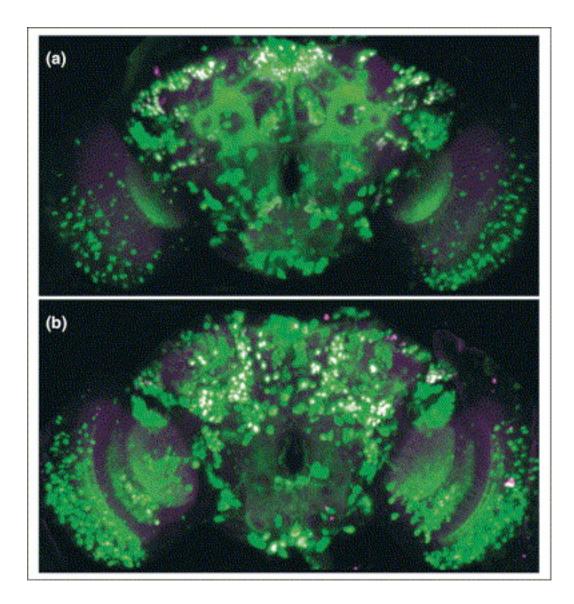


### males expressing female Fru push

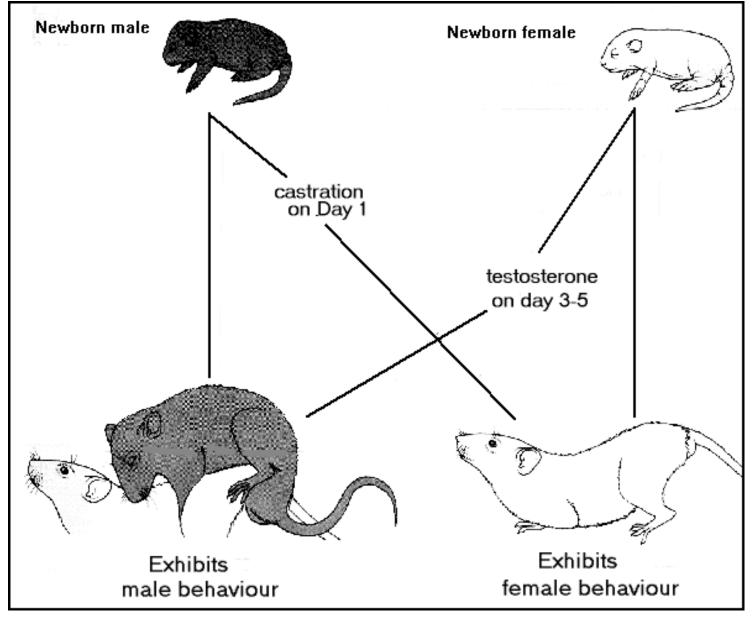
## Dominance relationships are also regulated by Fruitless

|                                | Probability of winning |                    |                   |                         |     |
|--------------------------------|------------------------|--------------------|-------------------|-------------------------|-----|
| Pairing                        | Overall                | As prior<br>winner | As prior<br>loser | Dominance<br>index (DI) | п   |
| <i>fru<sup>C</sup></i> males   | (0.50)                 | 0.88               | 0.12              | 0.75                    | 306 |
| <i>fru<sup>C</sup></i> females | (0.50)                 | 0.61               | 0.39              | 0.22                    | 156 |
| fru <sup>F</sup> males         | (0.50)                 | 0.69               | 0.31              | 0.39                    | 197 |

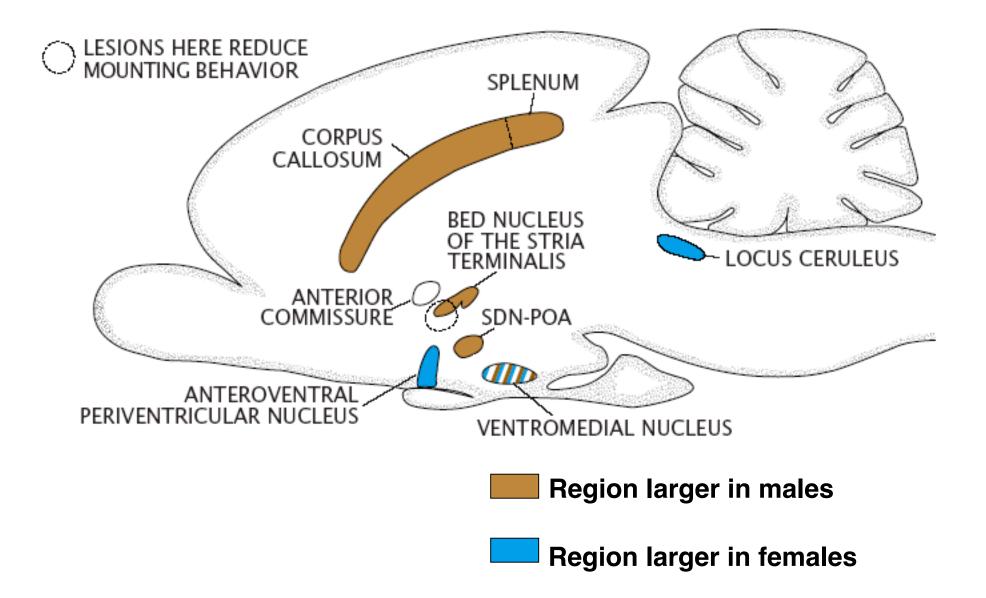
#### Fruitless is expressed in a subset of CNS neurons



# During early development testosterone regulates mating behavior in rats



### Sexually dimorphic regions of the rat brain



## **Sexual orientation in sheep**



## 8% of rams are male oriented; *i.e.,* they attempt to mate with rams. Male-oriented rams have a sexually dimorphic part of the brain that is similar in size to that in females.

#### The sheep SDN is sexually dimorphic

