

Model Organism

Model organisms for developmental biology research

Definition:

A model organism is a species that is extensively studied to understand particular biological phenomena.

Purpose:

General purpose:

provide insight into the workings of other organisms.

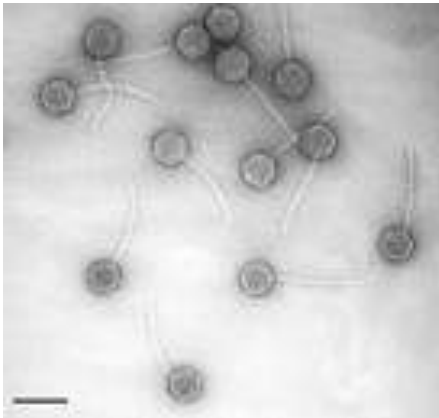
Specific purpose:

investigate the causes and treatments for human diseases when human experimentation would be unfeasible or unethical.

Model Organisms

1. Viruses: Phage Lambda 噬菌体
2. Prokaryotes: E. coli 大肠杆菌
3. Eukaryotes: Yeast 酵母菌
4. Plants: Arabidopsis 拟南芥

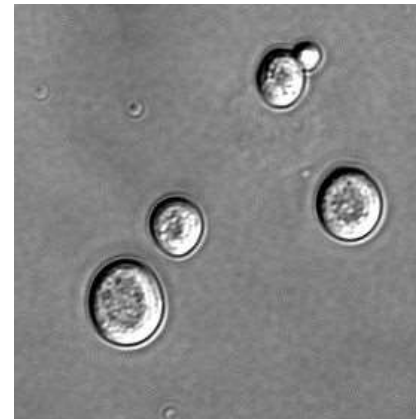
Lambda



E. Coli



S. cerevisiae



Arabidopsis



Animal Model Organisms

Invertebrates

sea urchin 海胆

C. elegans 线虫

D. melanogaster (fruit fly) 果蝇

sea urchin



C. elegans



Fruit fly



Animal Model Organisms

Vertebrates

zebrafish 斑马鱼

X. laevis (frog) 非洲爪蟾

chicken 鸡

mouse 小鼠

Zebrafish



X. laevis



chicken

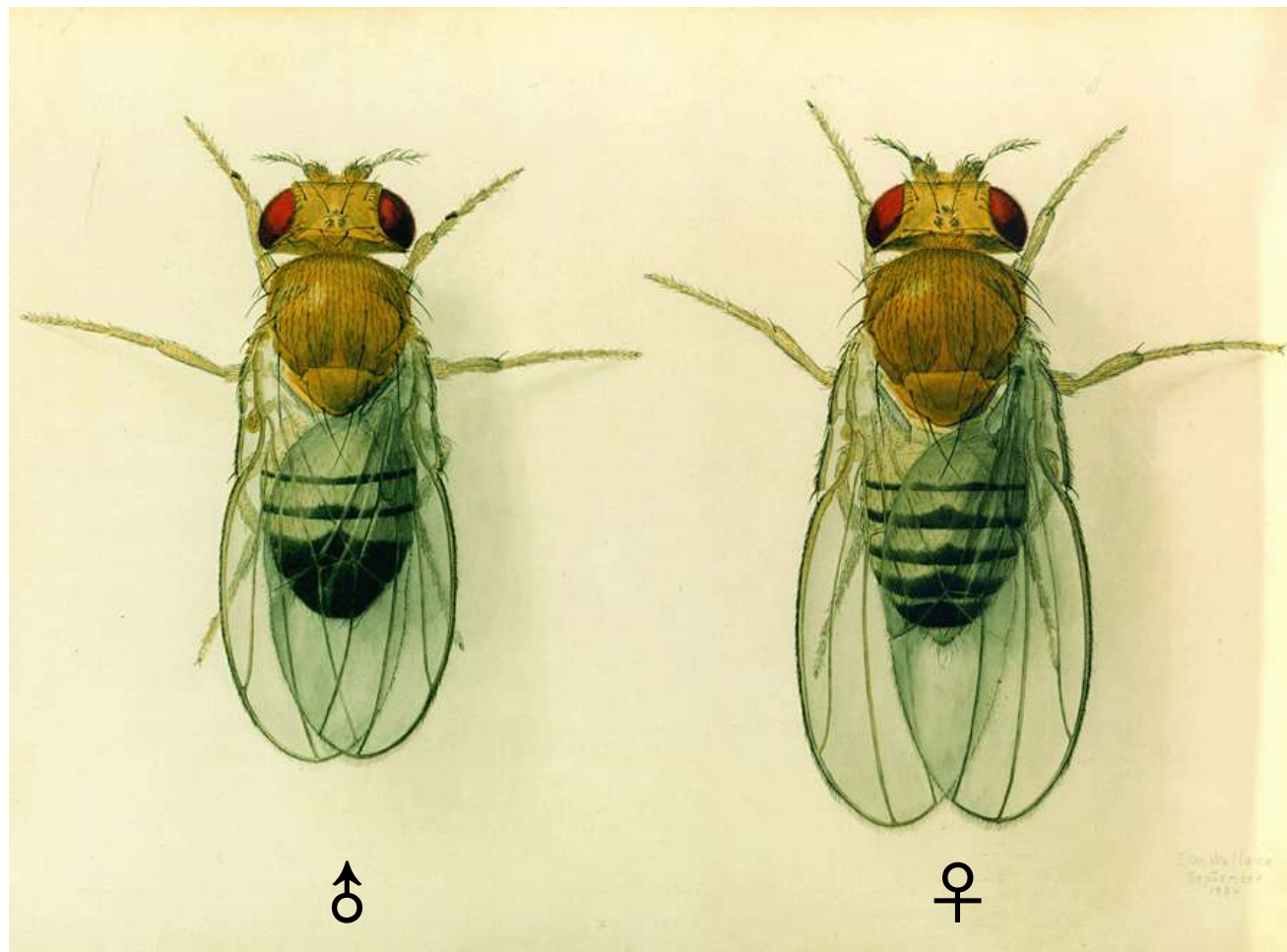


mouse



Drosophila melanogaster

黑腹果蝇



Drosophila 果蝇

means “dew-loving”

Latin: *drósos* *phílos*

English: dew loving
露水

also called: fruit fly

reason: linger around overripe or rotting
fruits, in particular, grapes.

Drosophila melanogaster

Scientific classification

<u>Kingdom:</u>	Animalia	动物界
Phylum:	Arthropoda	节肢动物门
Class:	Insecta	昆虫纲
Order:	Diptera	双翅目
Family:	Drosophilidae	果蝇科
Genus:	Drosophila	果蝇属
<u>Species:</u>	Drosophila melanogaster	黑腹果蝇种

Drosophila

about 1,500 species totally

found all around the world, in deserts, tropical rainforest, cities, and alpine zones

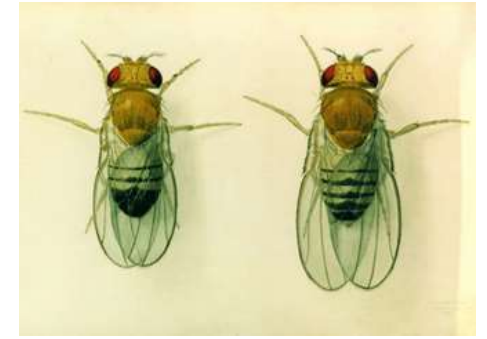
more species in the tropical regions

493 species in China

Food: fruit, decaying plant, flowers, fungal and mushrooms

Drosophila melanogaster

The most important model organism for studies in
Genetics and Developmental Biology



Small size:	5 mm	
A short generation time:	10 days	
Large number of offspring:	a female lays 50 - 80 eggs everyday	
Life span:	60 - 90 days	
Compact genome:	sex chromosome	1 pair
	autosome	3 pairs
Easy to obtain mutant animals:	X-ray, chemical mutagen (EMS),	
	transposable element	
Genetic manipulation is easy :	100 years of genetics	
Easily cultured and Inexpensive:	fruits or medium	
Genome:	sequenced in 2000, about 15000 genes	
	77% of human disease genes (714/929 genes)	



The Nobel Prize in Physiology or Medicine 1933

"for his discoveries concerning the role played by the chromosome in heredity "



Thomas Hunt Morgan

(1866 –1945)



1906, began his work on *D. melanogaster* at Columbia University

1910, reported the white eyed mutant – the first gene (white) identified

1933, rewarded Nobel Prize in Medicine

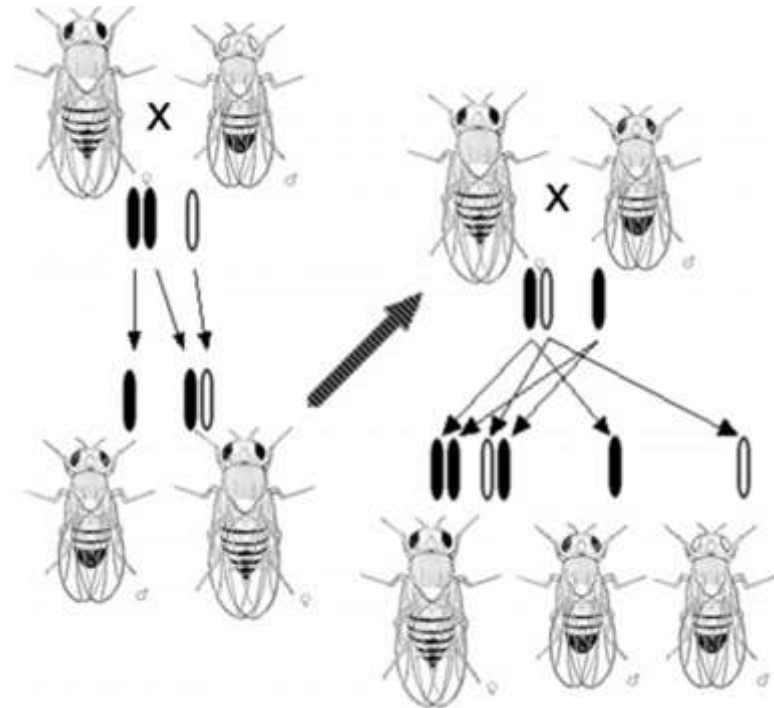
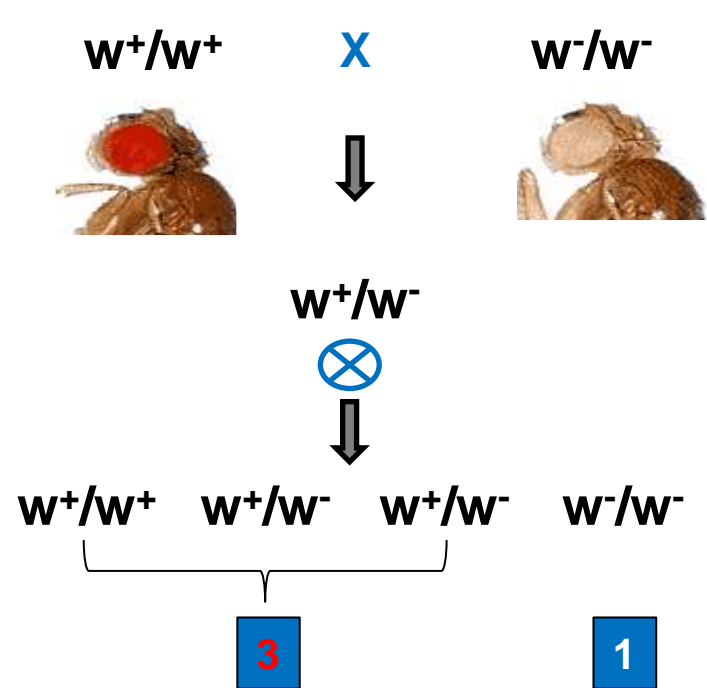
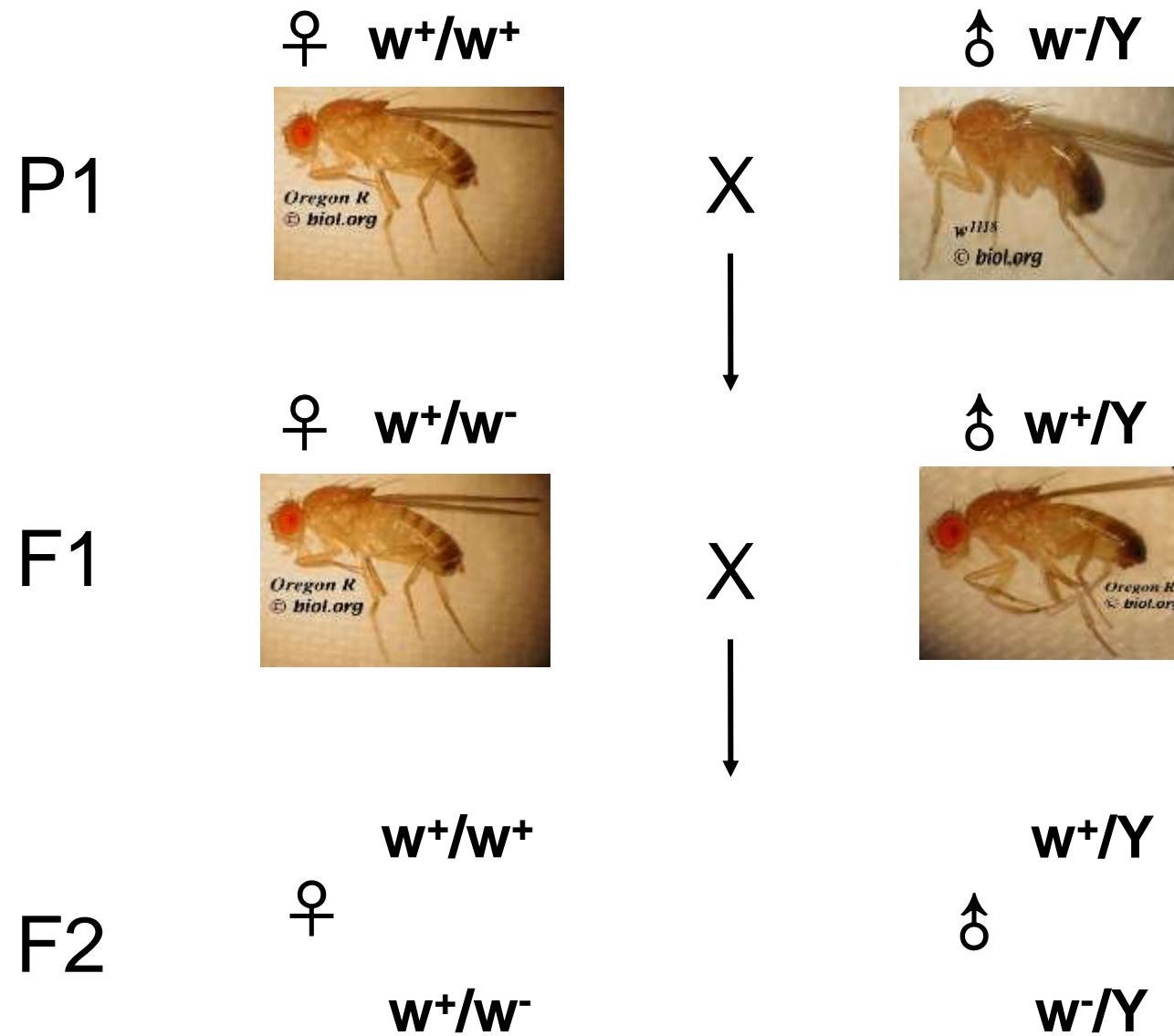
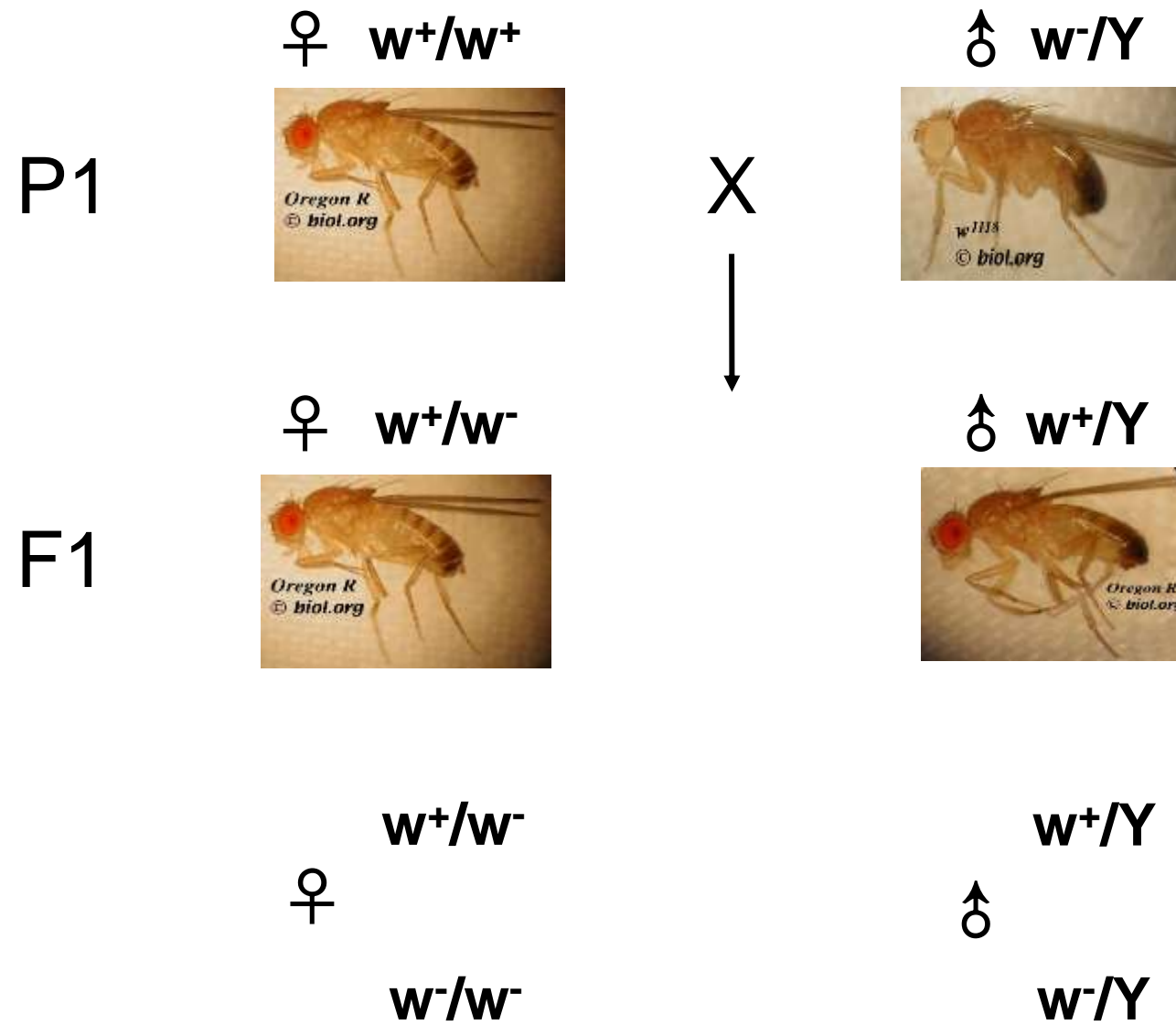


Table 1: Expected Mendelian Ratios versus Morgan's Actual Results

Cross	Outcome	
	Expected Phenotypes	Observed Phenotypes
P ₁ Red ♀ × P ₁ White ♂	F ₁ = All Red	F ₁ = All Red*







Thomas Morgan



Alfred Sturtevant

Identified chromosomes as the carriers of the hereditary material as a undergraduate student of Morgan in 1911

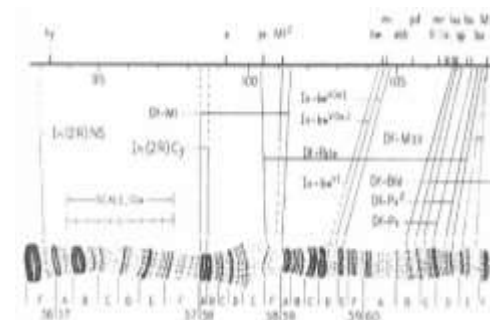


Calvin Bridges

Ph.D. thesis - the 1st paper in the 1st issue of *Genetics* in 1916



Polytene chromosome and banding pattern





Thomas Morgan

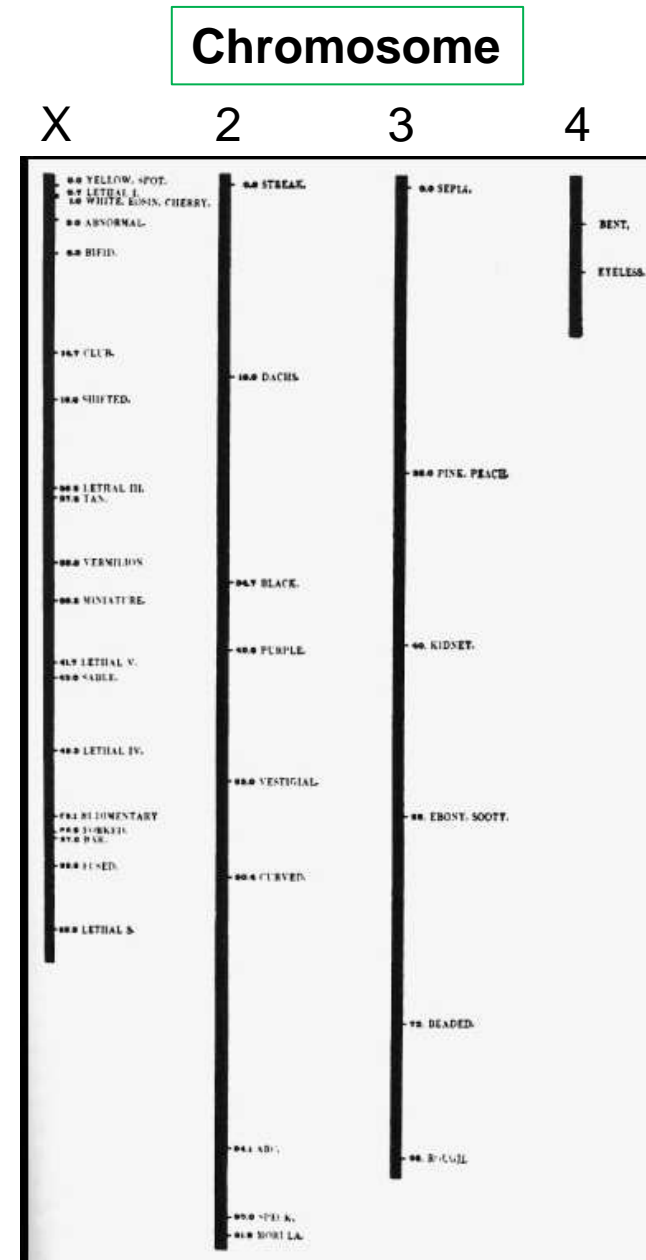


Alfred Sturtevant



Calvin Bridges

Ph.D. thesis - the 1st paper
in the 1st issue of *Genetics* in 1916





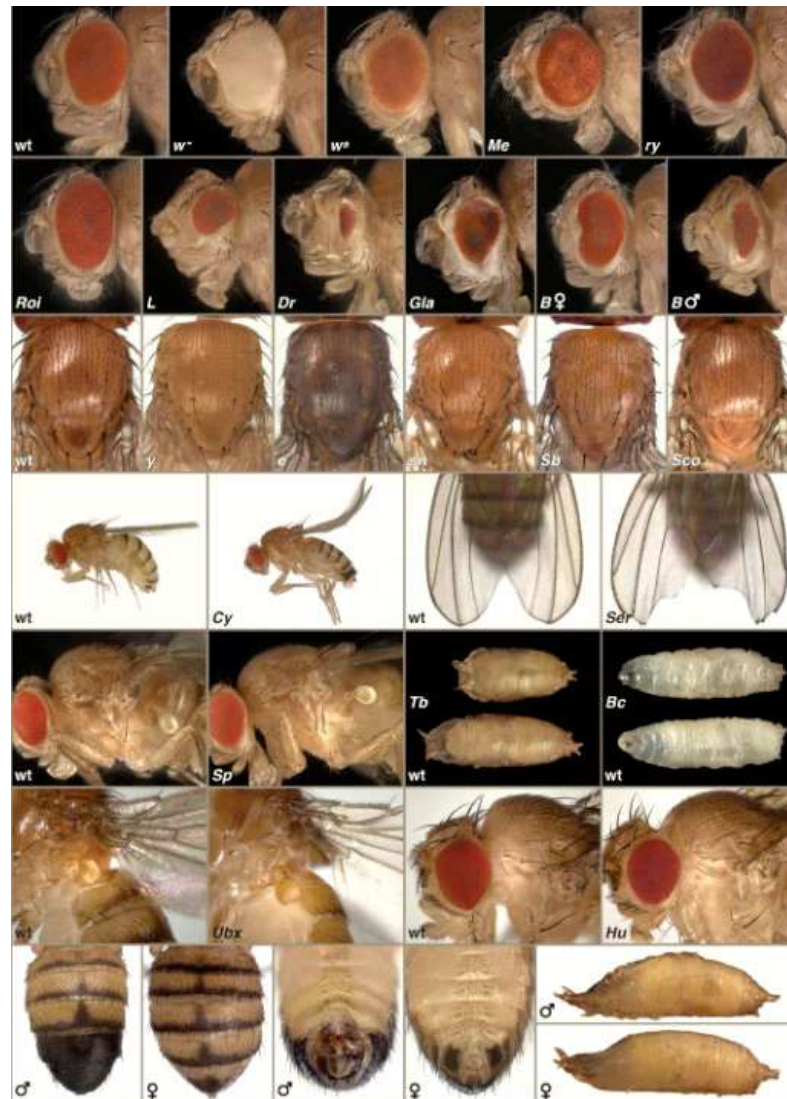
The Nobel Prize in Physiology or Medicine 1946

"for the discovery of the production of mutations by means of X-ray irradiation "

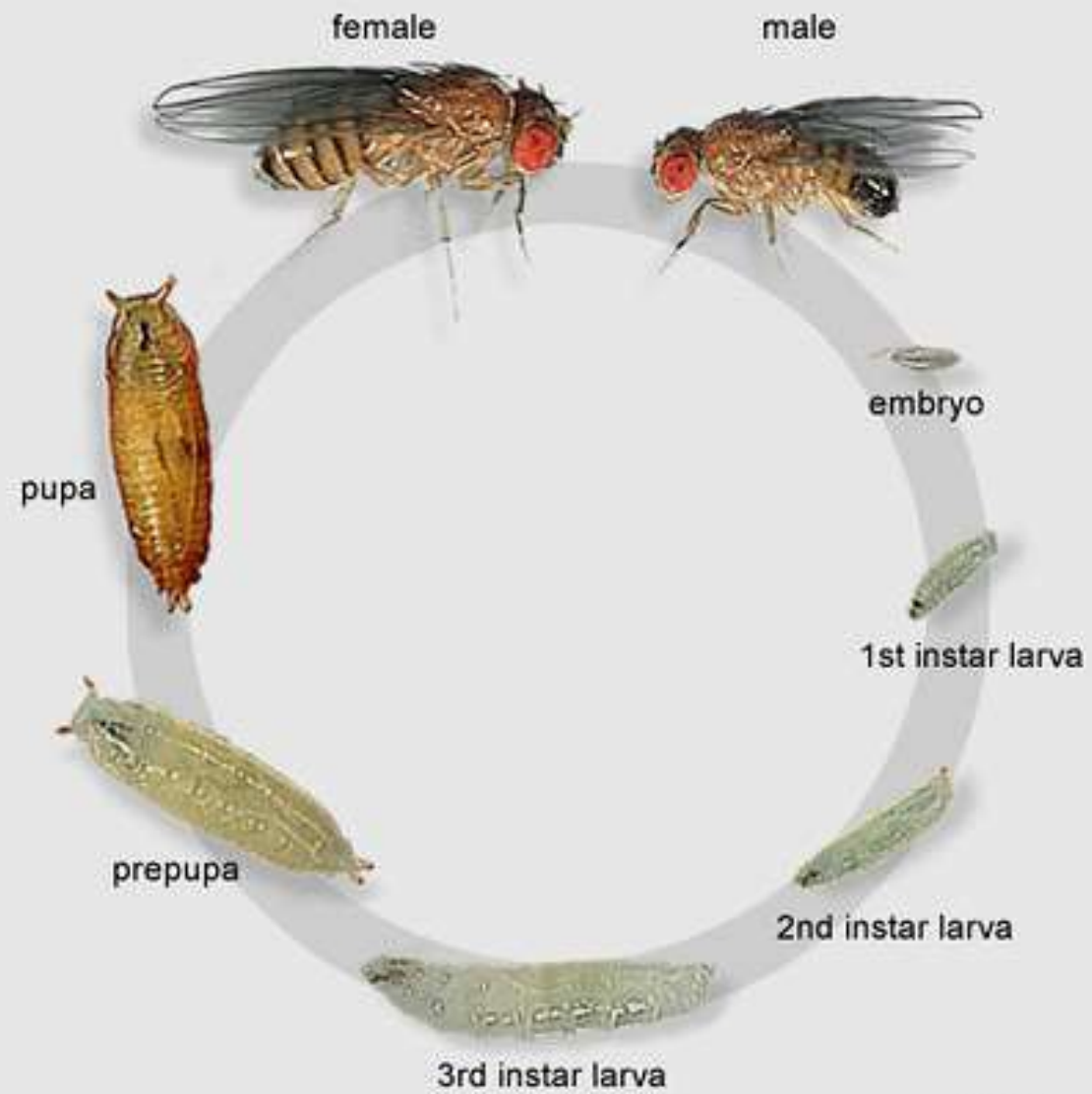


Hermann Joseph Muller

(1890 –1967)



The life cycle of *Drosophila melanogaster*





Drosophila oocyte

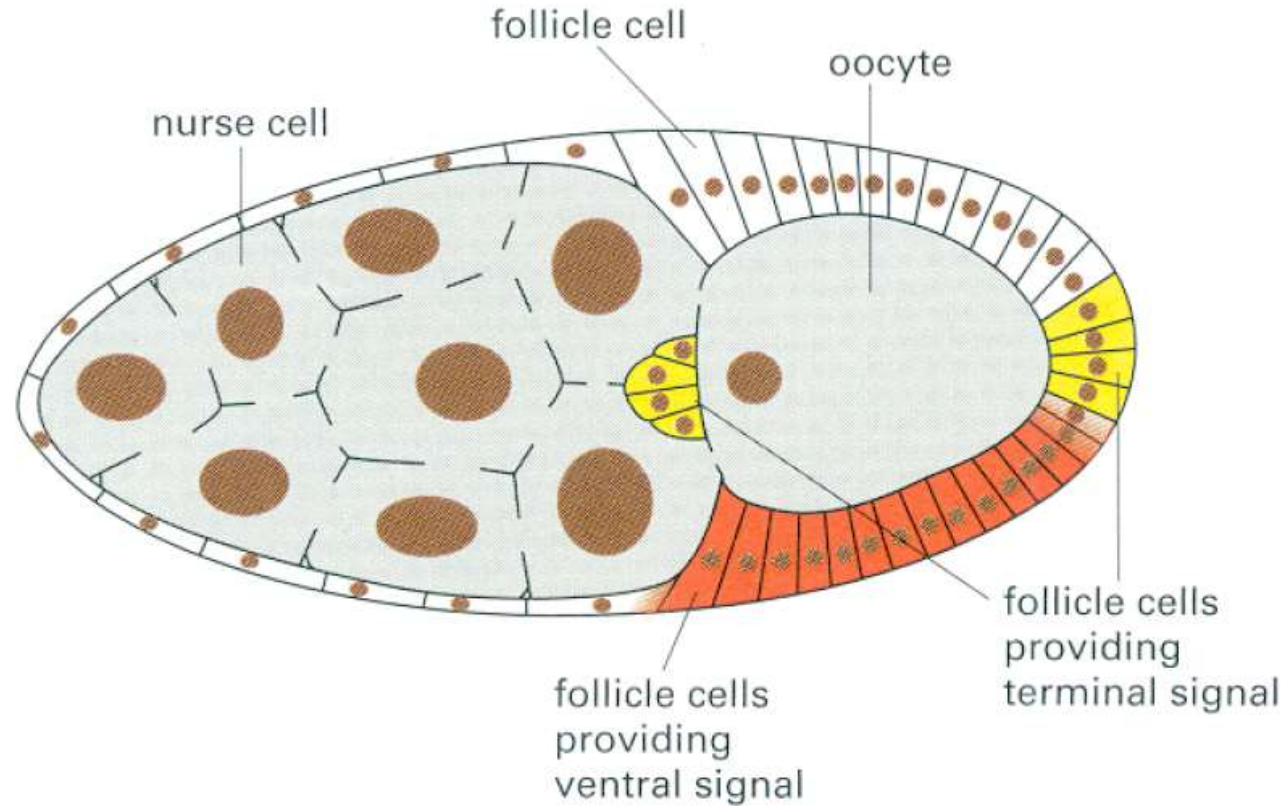
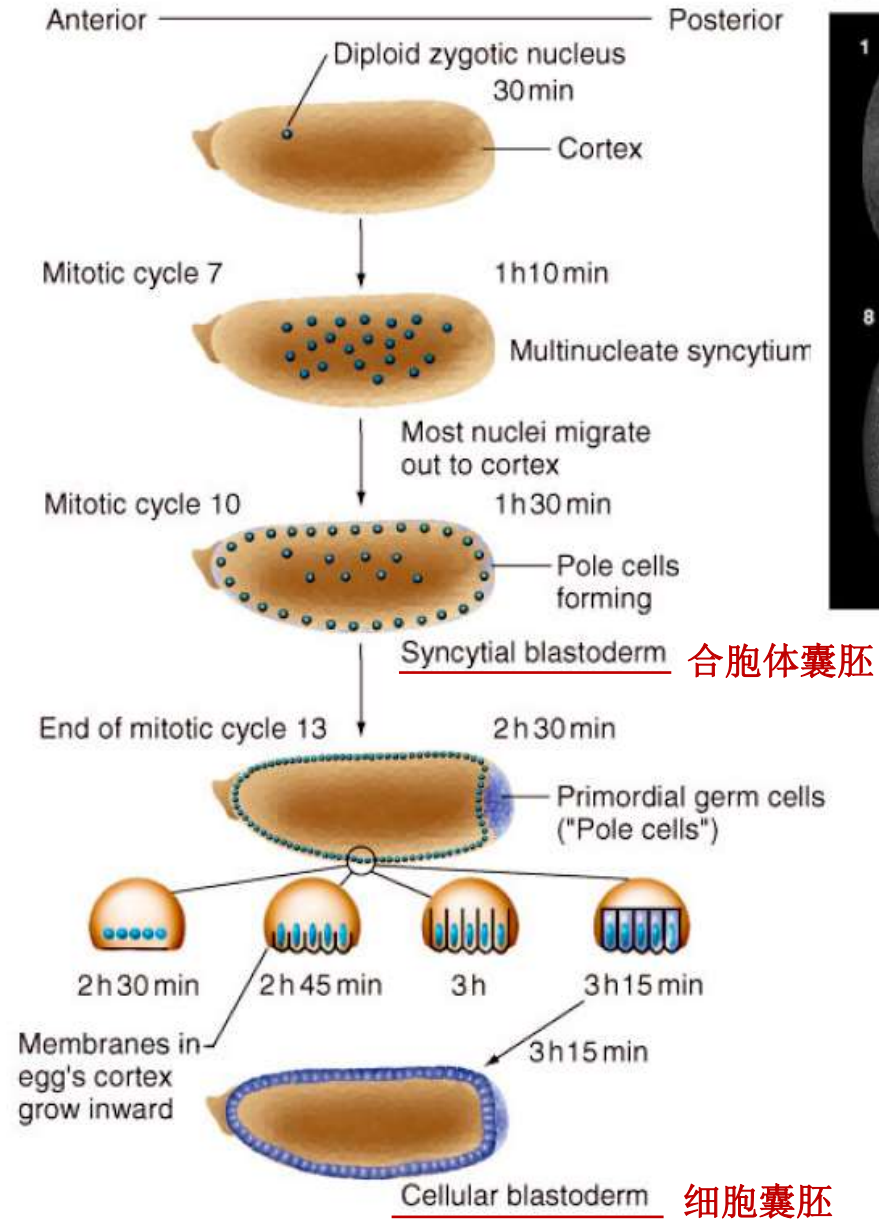
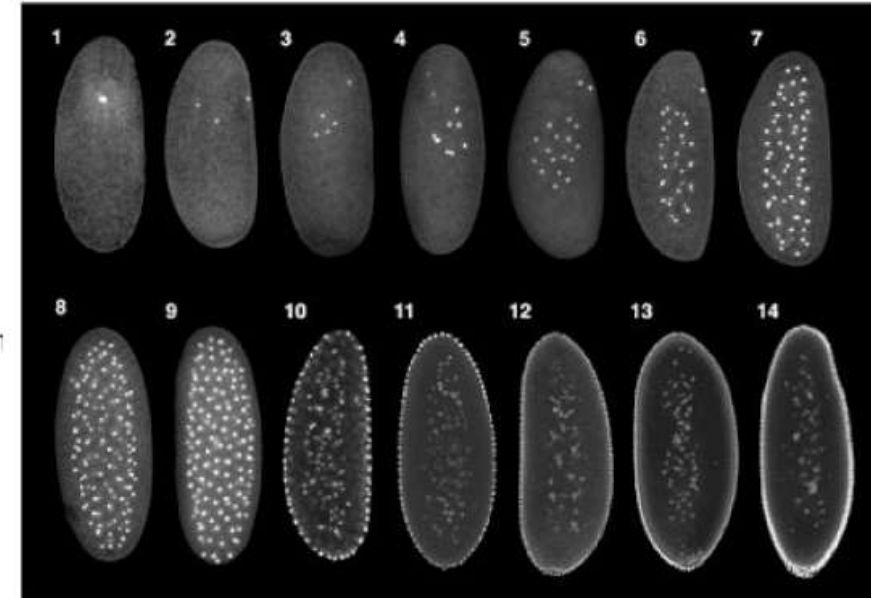


Figure 21-30. A *Drosophila* oocyte in its follicle. The oocyte is derived from a germ cell that divides four times to give a family of 16 cells that remain in communication with one another via cytoplasmic bridges (gray). One member of the family group becomes the oocyte, while the others become nurse cells, which make many of the components required by the oocyte and pass them into it via the cytoplasmic bridges. The follicle cells that partially surround the oocyte have a separate ancestry. As indicated, they are the sources of terminal and ventral egg-polarizing signals. (From **Bruce Albert Book**)

(a) The first three hours after fertilization

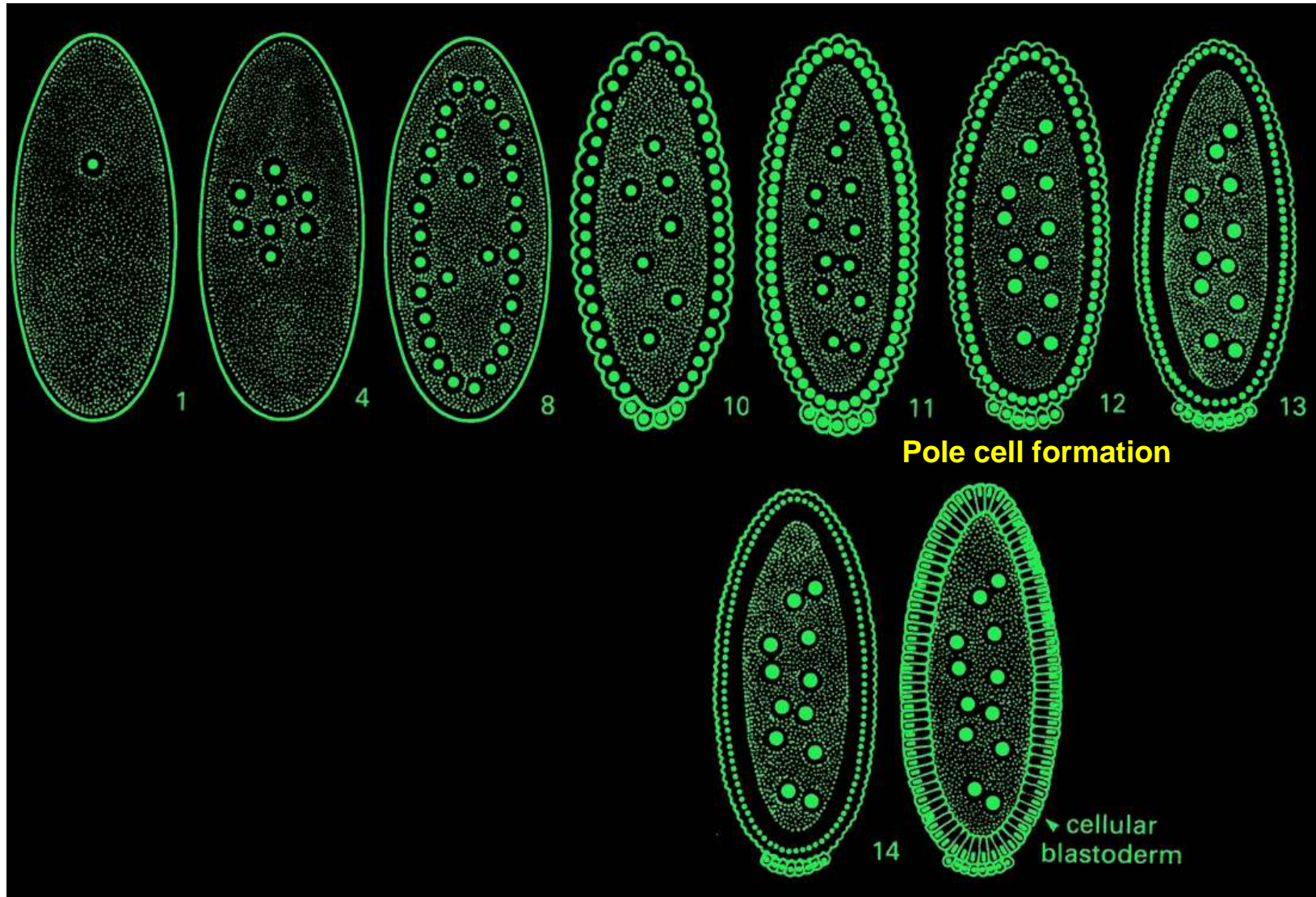


(b) Early embryonic stages in cross section



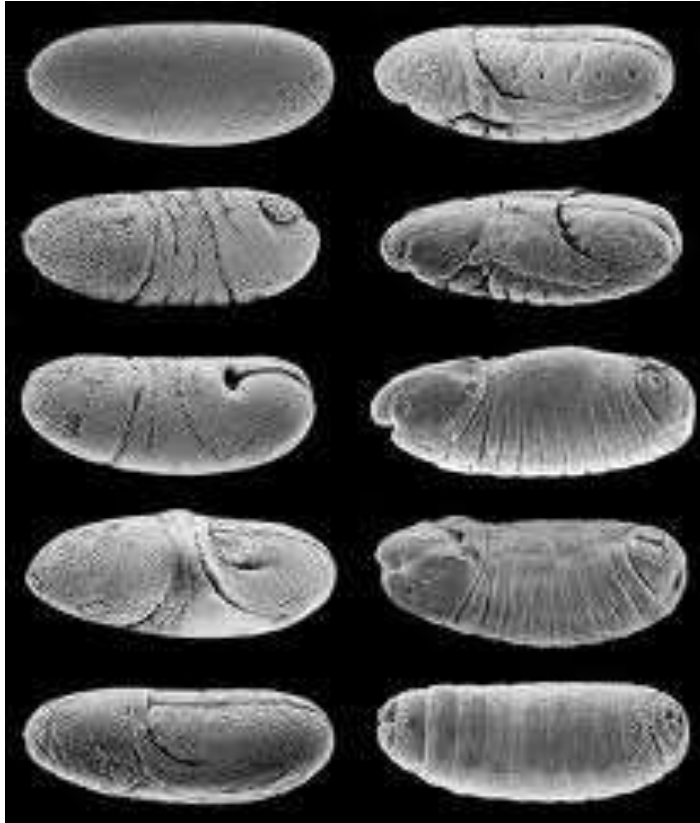
Drosophila Embryogenesis

syncytial blastoderm



Drosophila embryogenesis

Dorsal



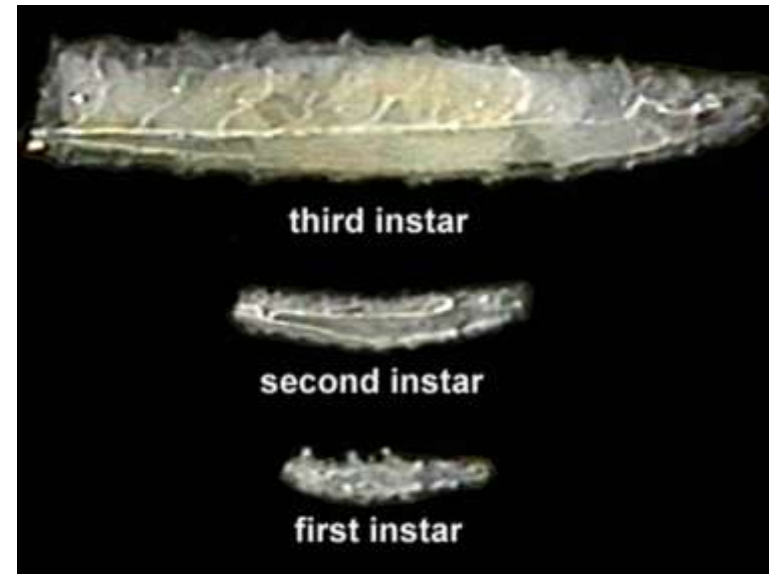
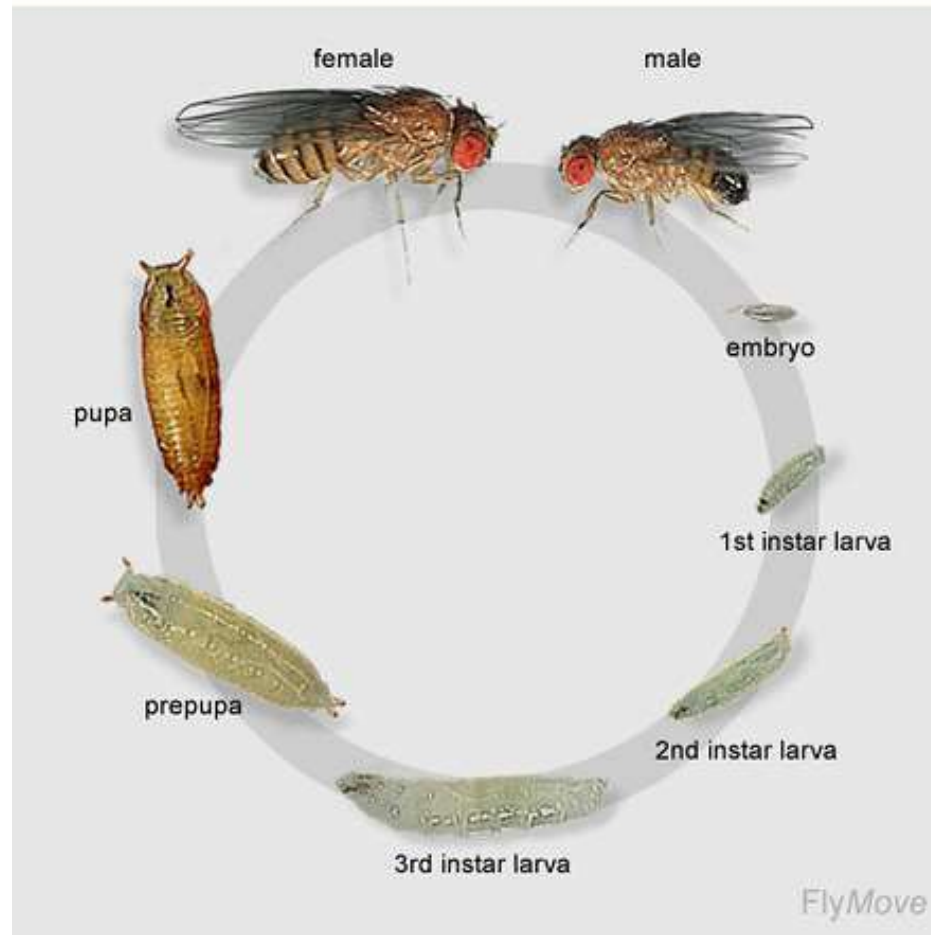
Ventral

Drosophila embryogenesis

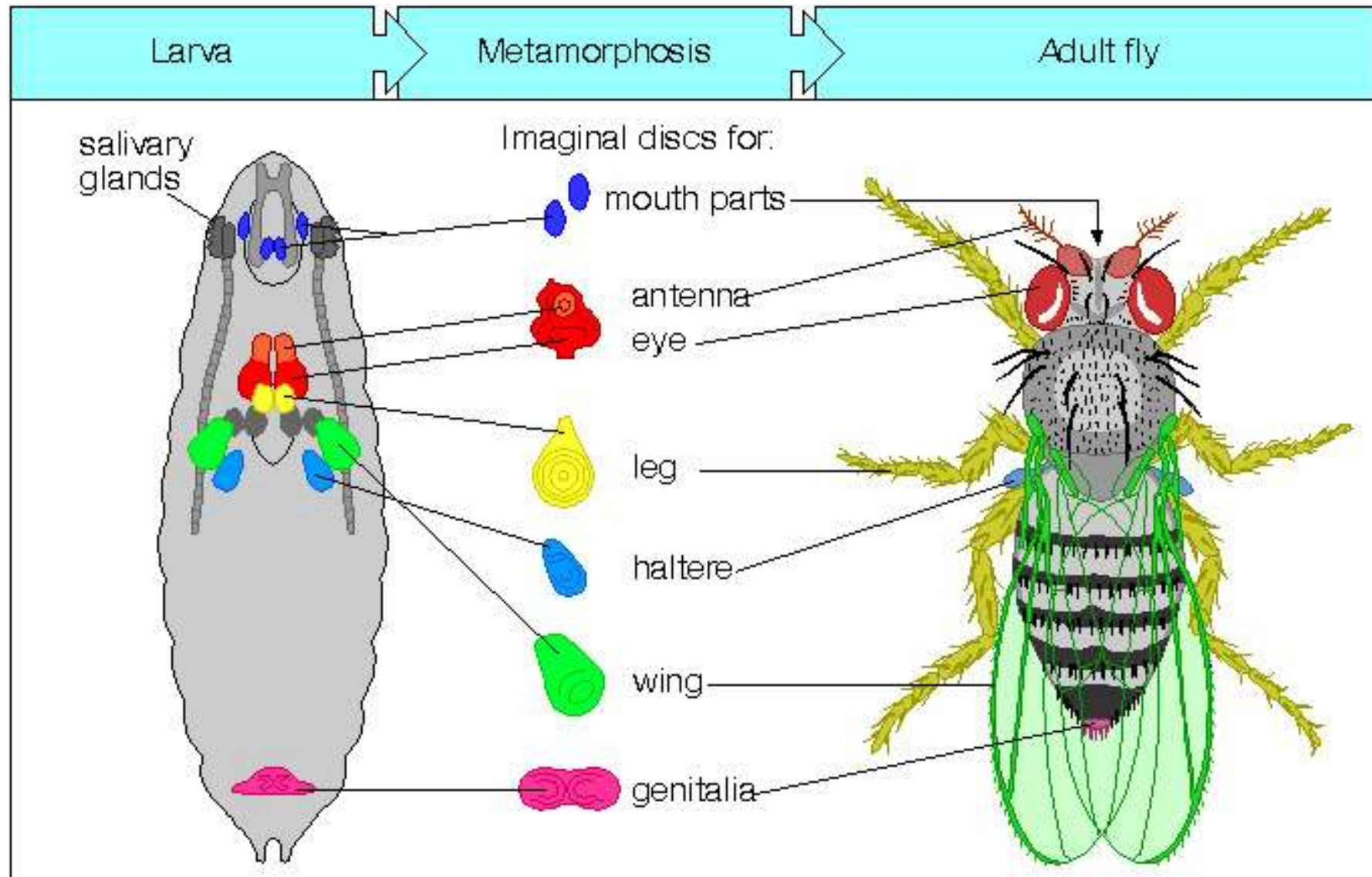
Drosophila embryo morphogenesis is the process from which a fertilized single cell egg becomes a multicellular embryo with structured tissues and specialized cells/organs.

This process occurs through many complex cell shape changes and movements, including the formation of cellular blastoderm, gastrulation, and germ band elongation/retraction.

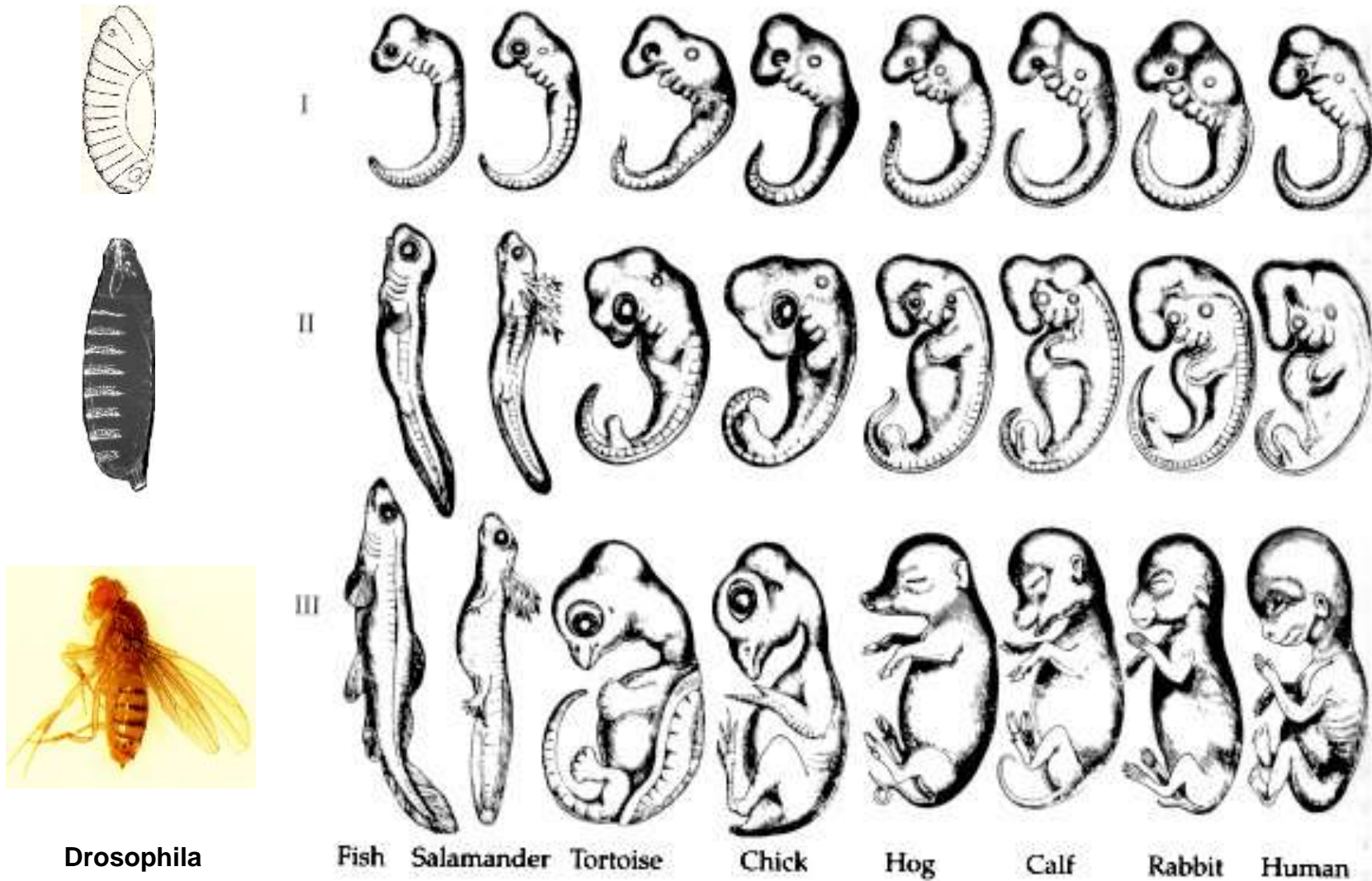
The life cycle of *Drosophila melanogaster*



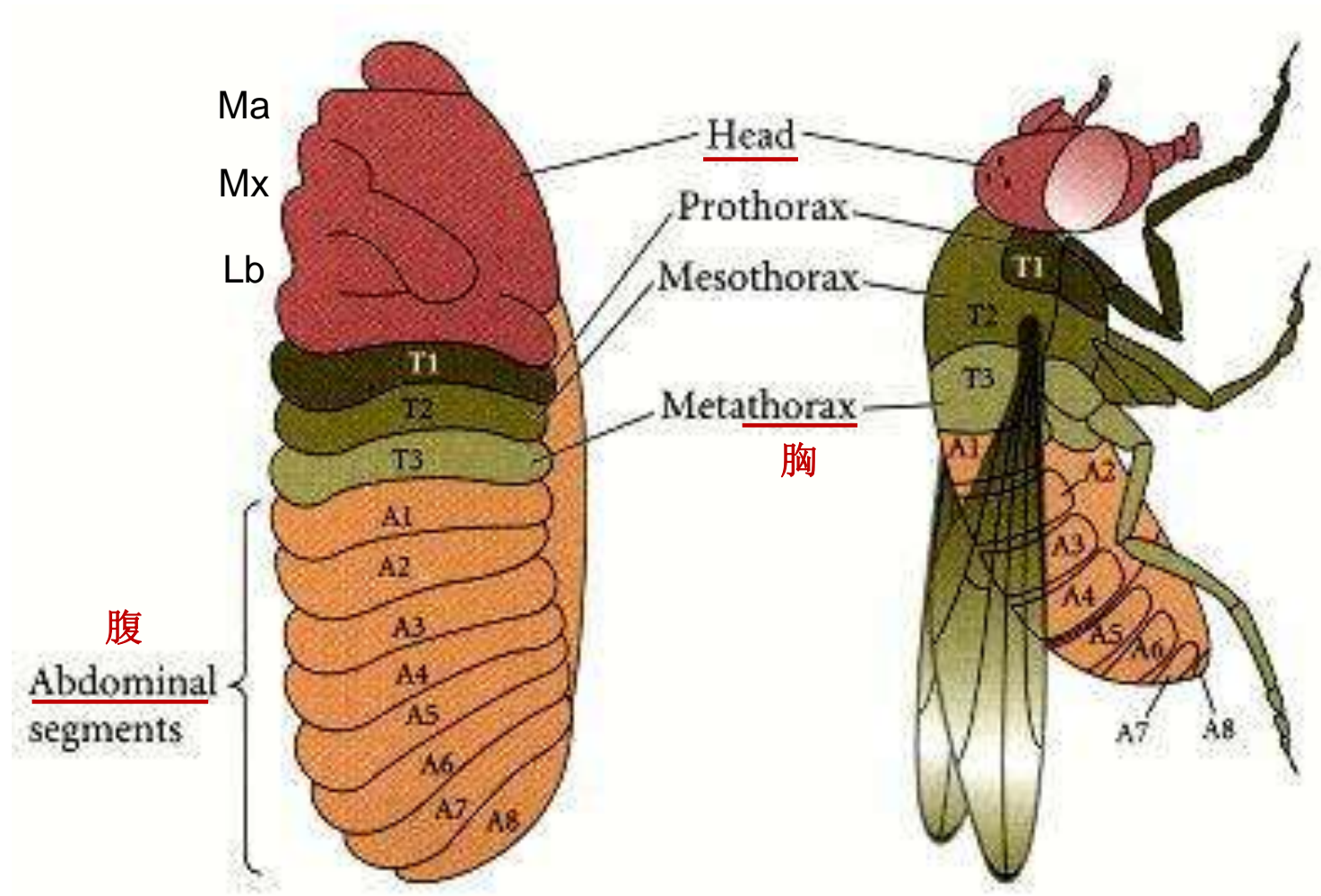
Drosophila Development



Haeckel's 1874 version of vertebrate embryonic development.



Segmentation in *Drosophila*





The Nobel Prize in Physiology or Medicine 1995

"for their discoveries concerning the genetic control of early embryonic development "



Edward B. Lewis



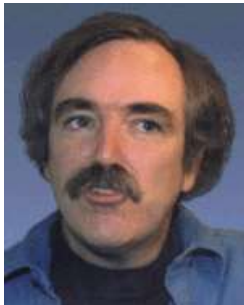
**Christiane
Nüsslein-Volhard**



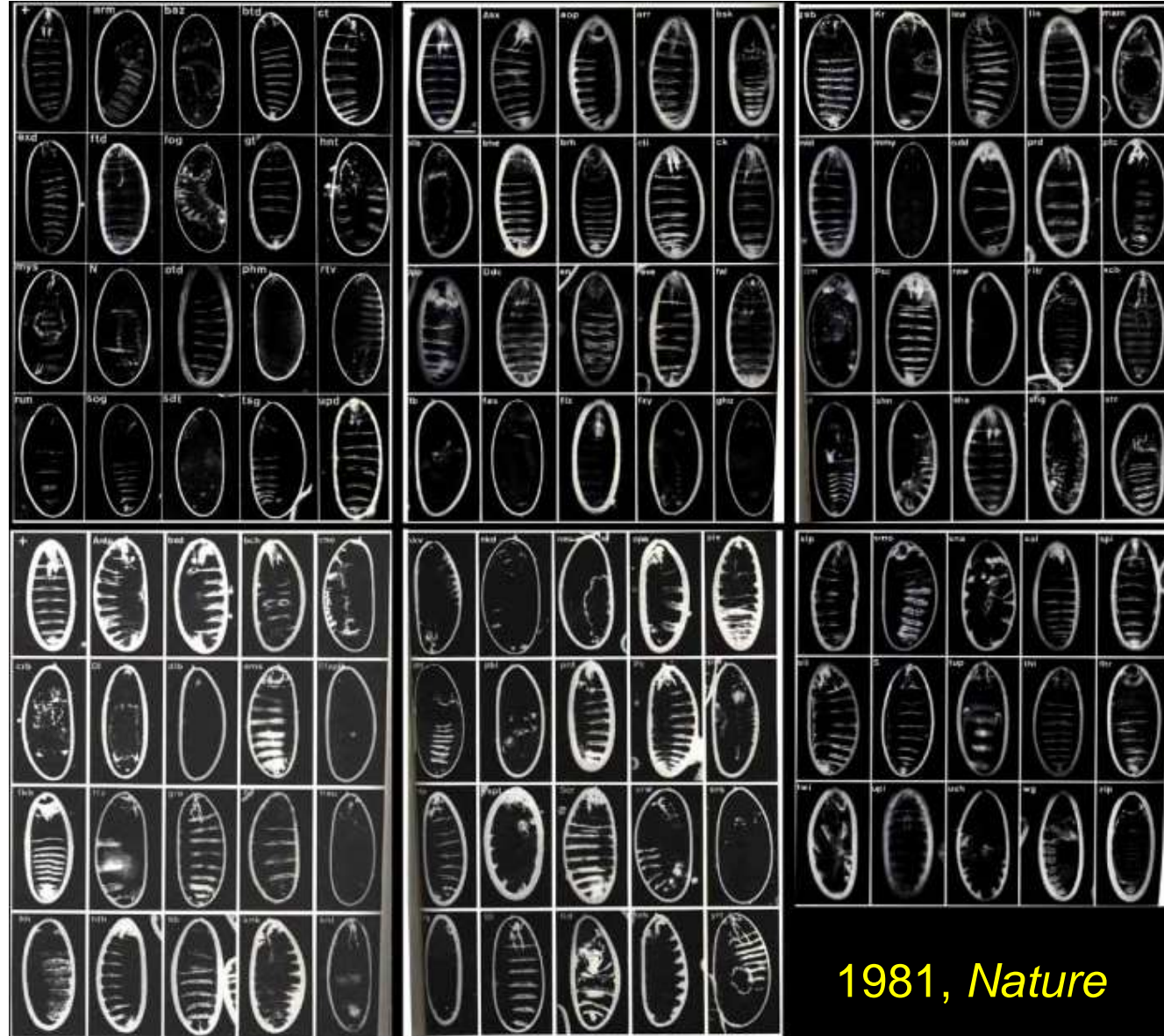
Eric Wieschaus



**Christiane
Nüsslein-Volhard**

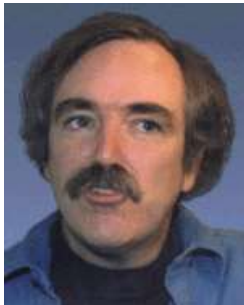


Eric Wieschaus





Christiane
Nüsslein-Volhard



Eric Wieschaus

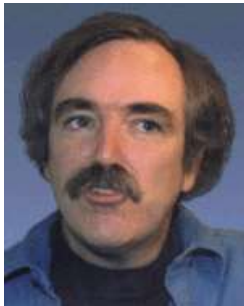
Four classes of genes responsible for formation of segments

- Maternal genes
- Gap genes
- Pair-rule genes
- Segmentation polarity genes

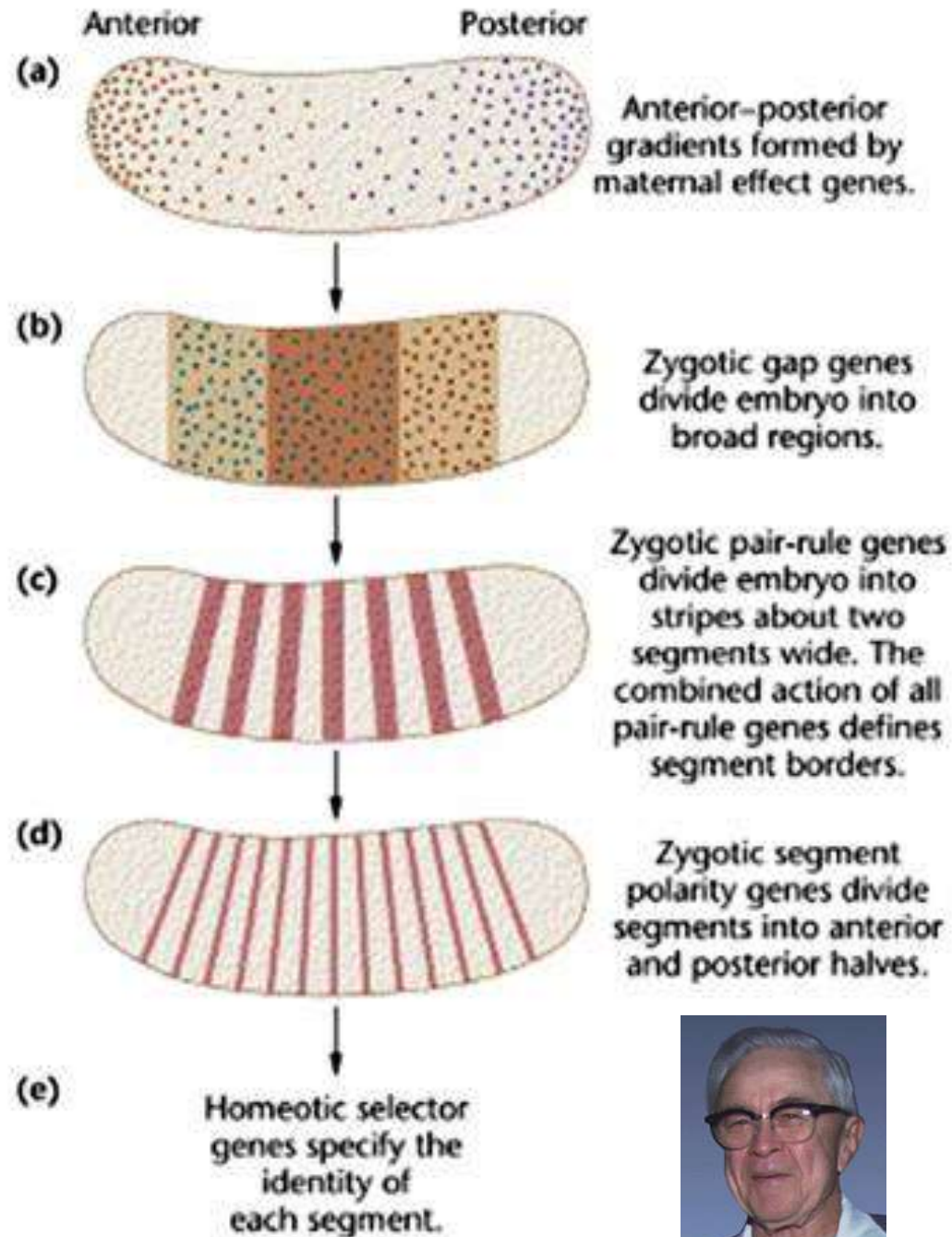
Function in hierarchy
that progressively subdivides
the embryo into successively small units



Christiane
Nüsslein-Volhard

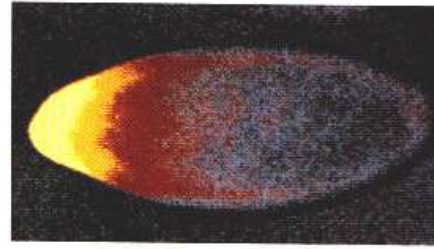
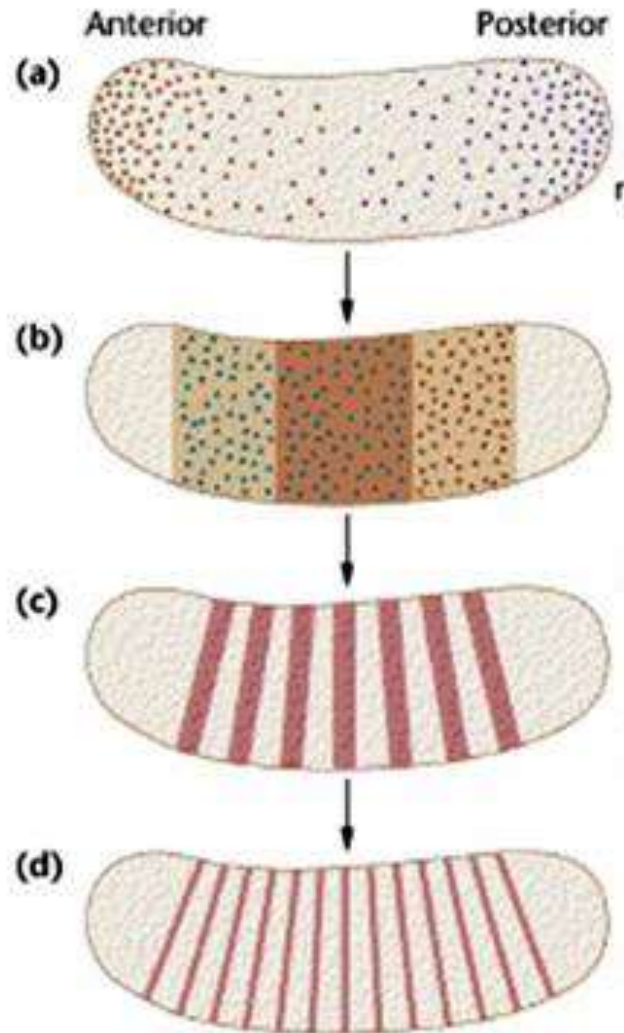


Eric Wieschaus



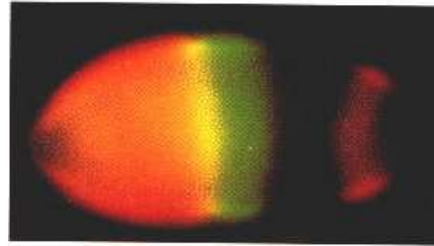
Edward B. Lewis

Anterior-posterior (A/P) polarity and Segmentation



bicoid

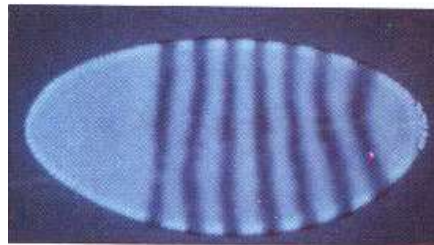
Maternal gene



Hunchback (red)

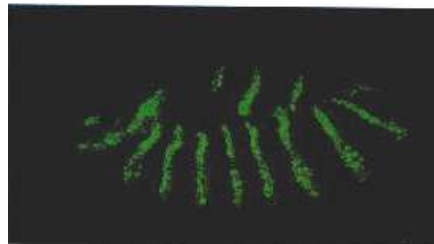
Kruppel (green)

Gap gene



fushi tarazu

Pair-rule gene



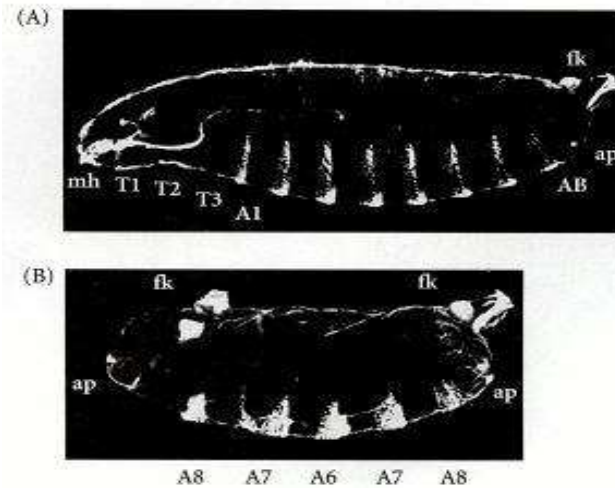
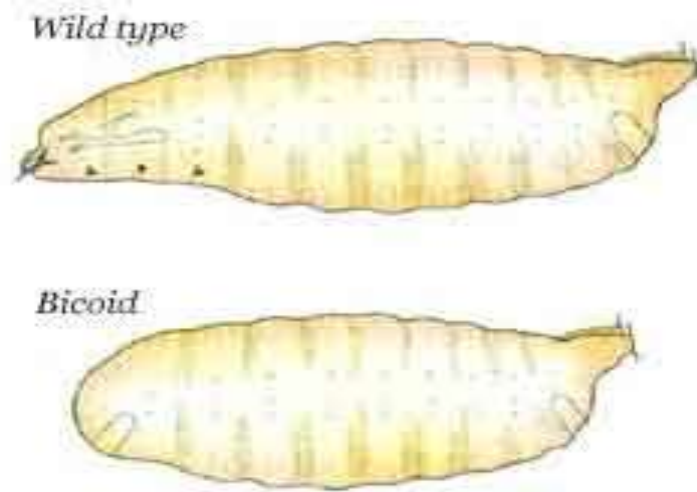
engrailed

Segment polarity gene

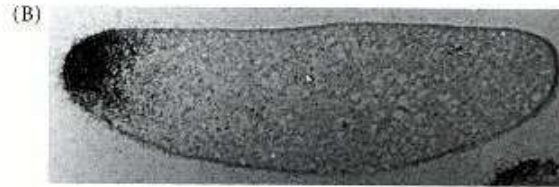


Eric Wieschaus

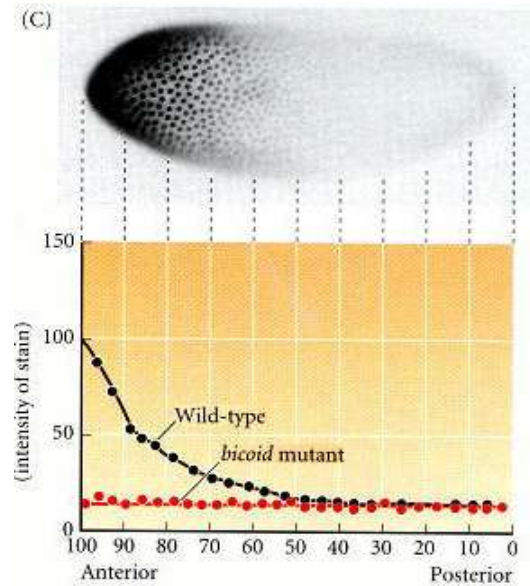
Maternal gene *bicoid* control anterior structures



bicoid is a **morphogen** 形态发生素



bicoid mRNA



bicoid protein

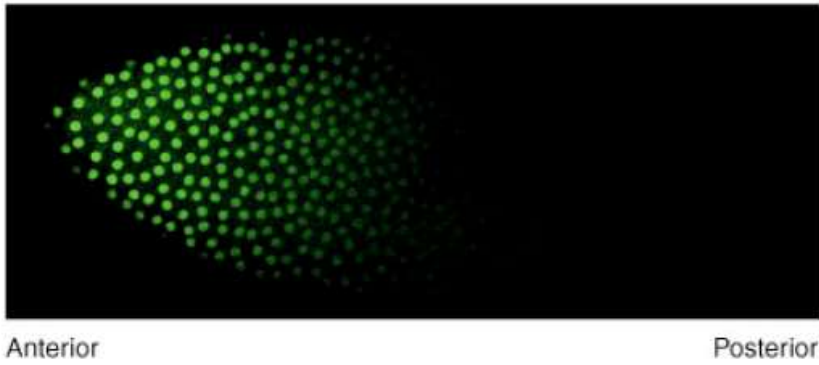
Morphogens: Substances that define different cell fate in a concentration-dependent manner

bicoid is a morphogen 形态发生素

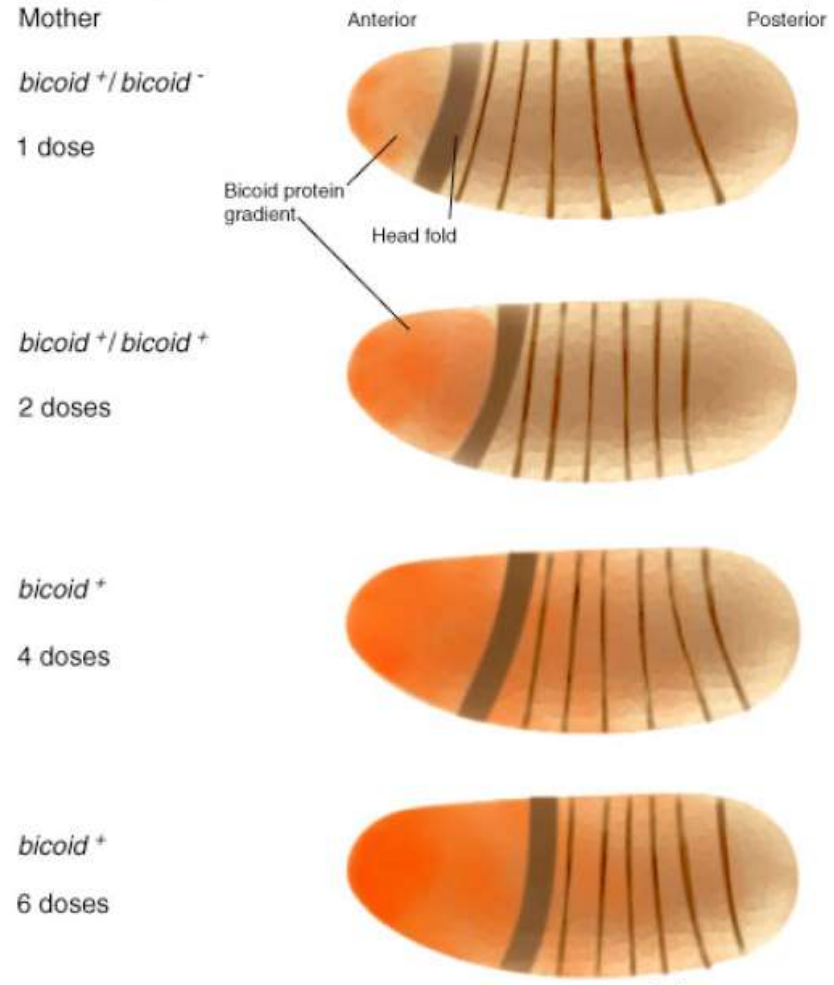
(a) Localization of *bicoid* mRNA



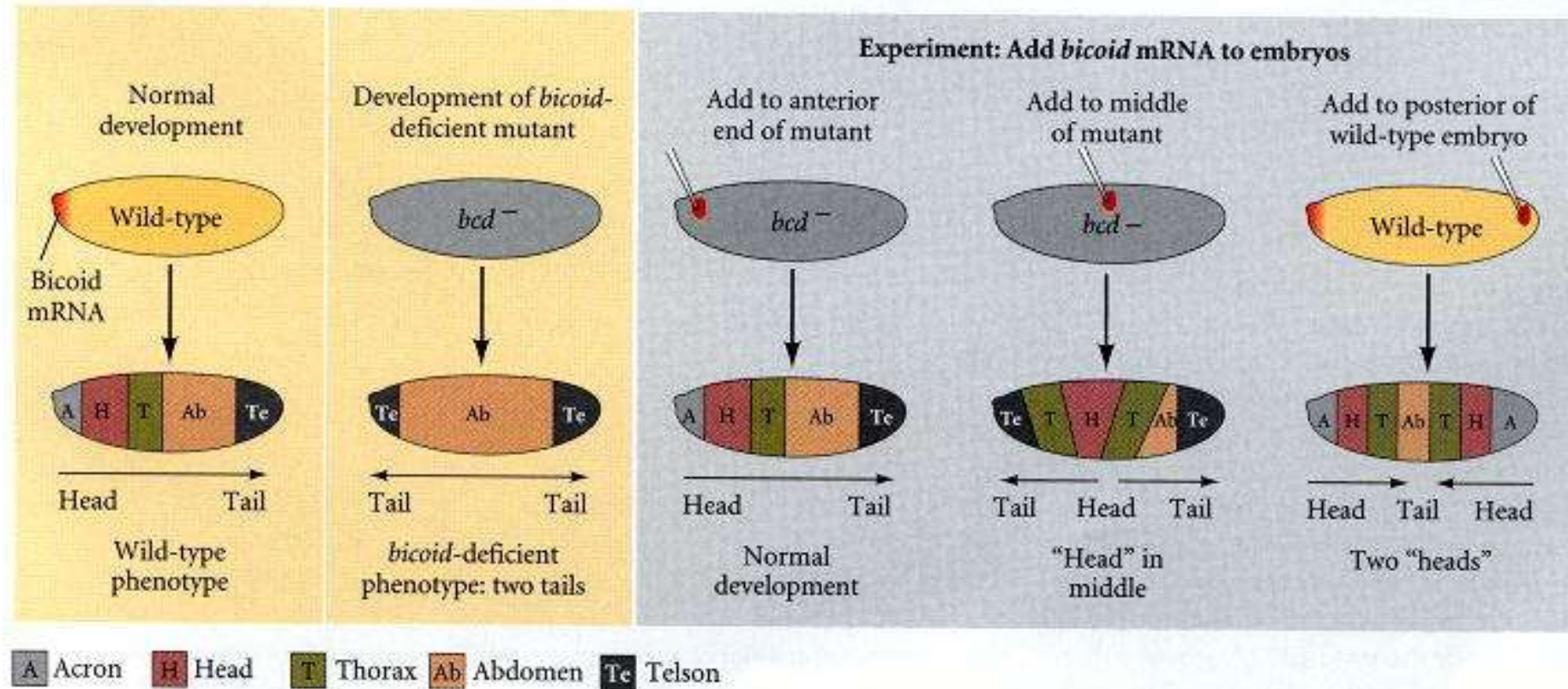
(b) A gradient of Bicoid protein



(c) Bicoid protein is a morphogen.



Maternal gene *bicoid* control anterior structures

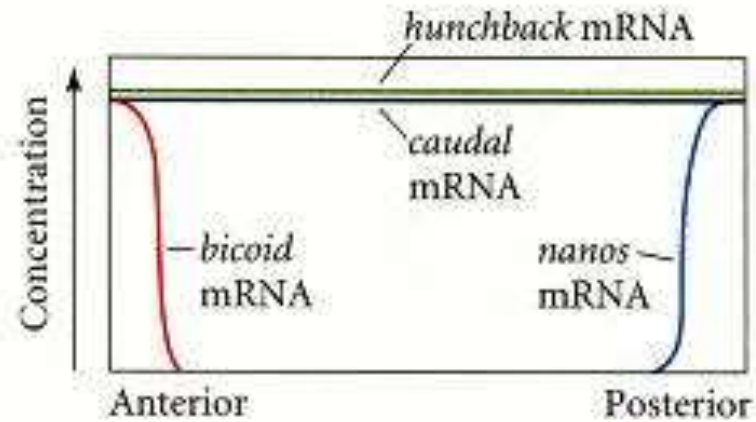


顶节

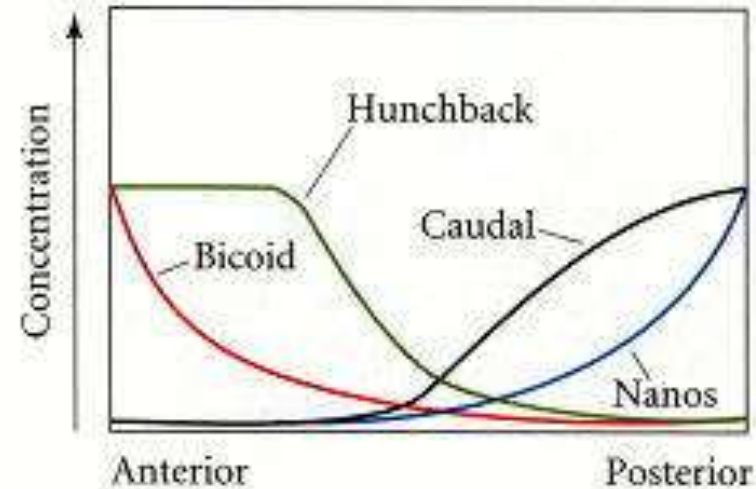
尾节

Maternal Genes

(A) Oocyte mRNAs

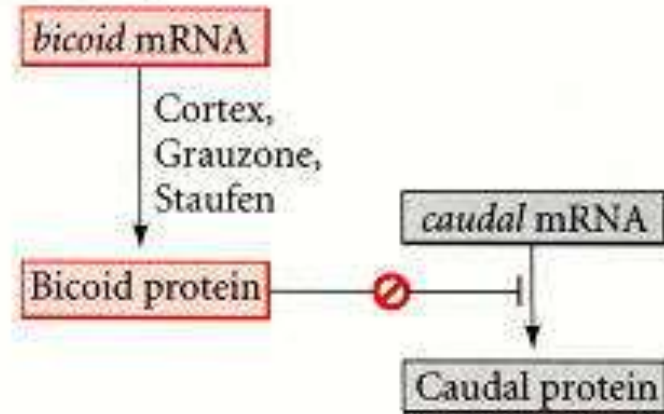


(B) Early cleavage embryo proteins

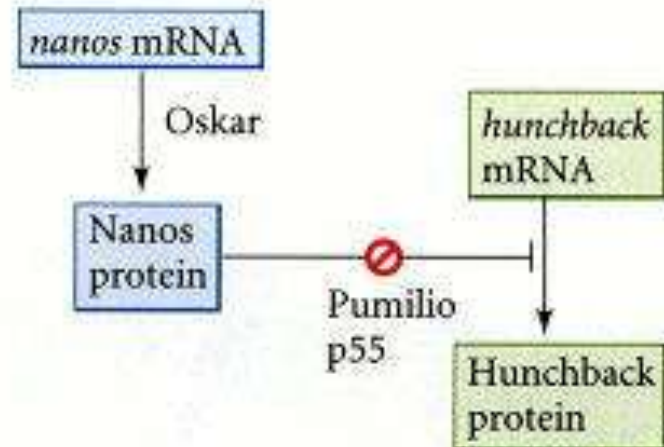


(C)

ANTERIOR



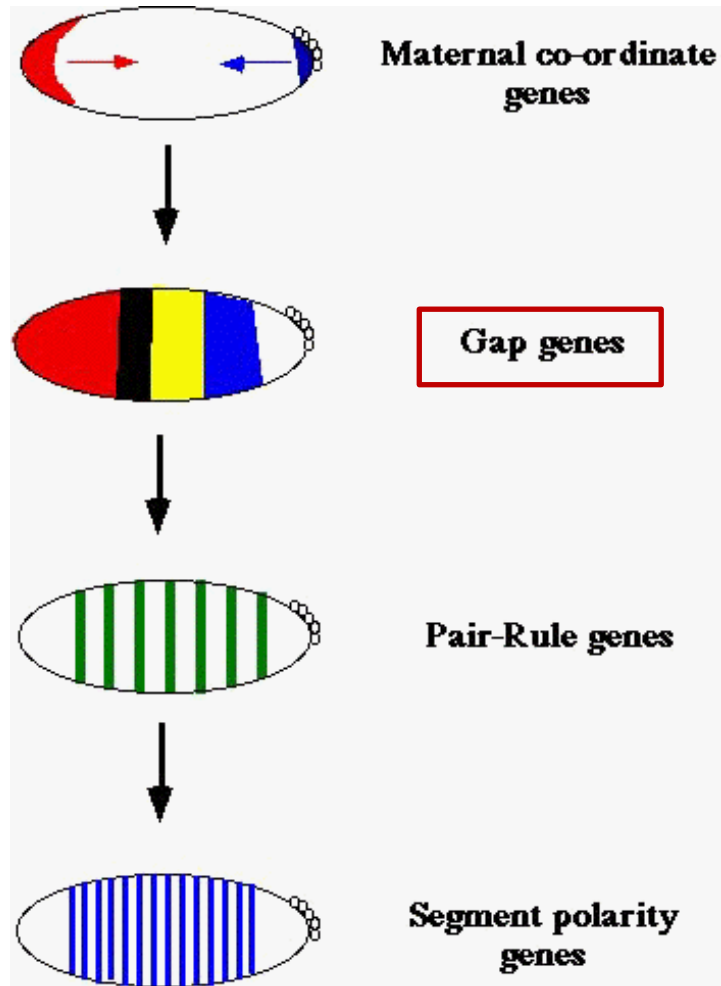
POSTERIOR





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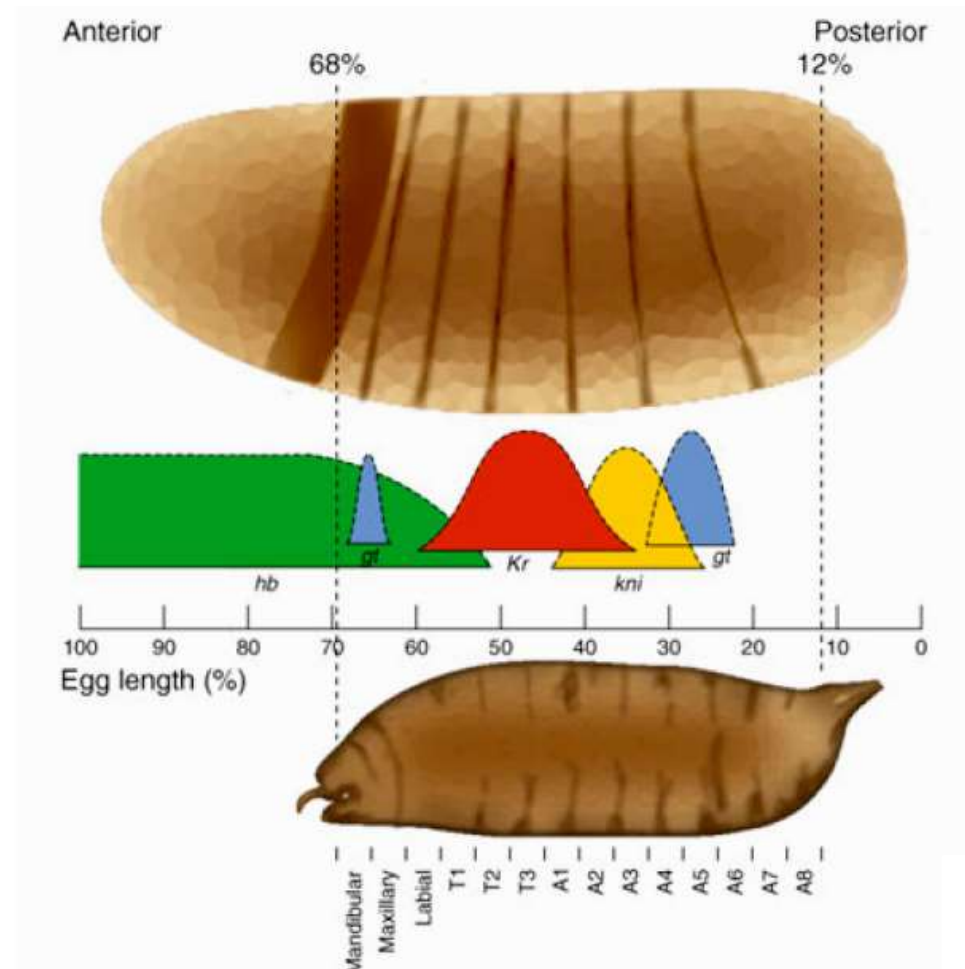
Anterior-posterior (A/P) polarity and Segmentation



Gap genes

- Gap mutants show a gap in segmentation pattern at positions where particular gene is absent
- Binding sites in promoter have different affinities for maternal transcription factors
- Gap genes encode transcription factors that influence expression of other gap genes

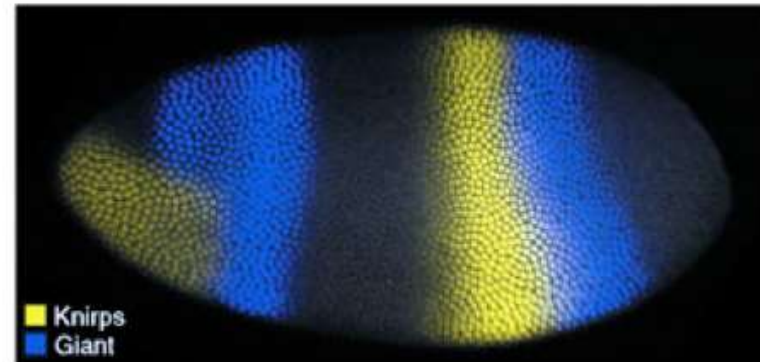
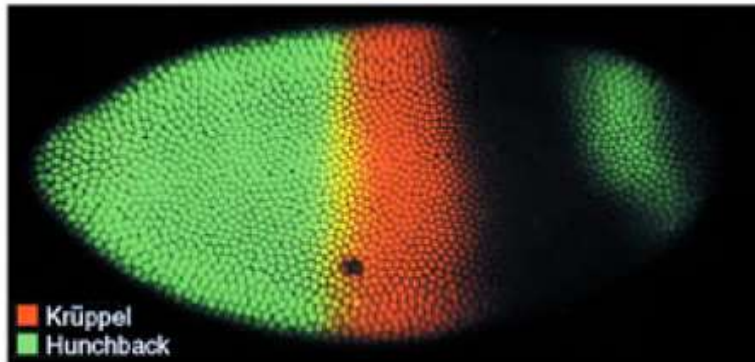
Mutations in gap gene result in loss of segments corresponding to zone of expression



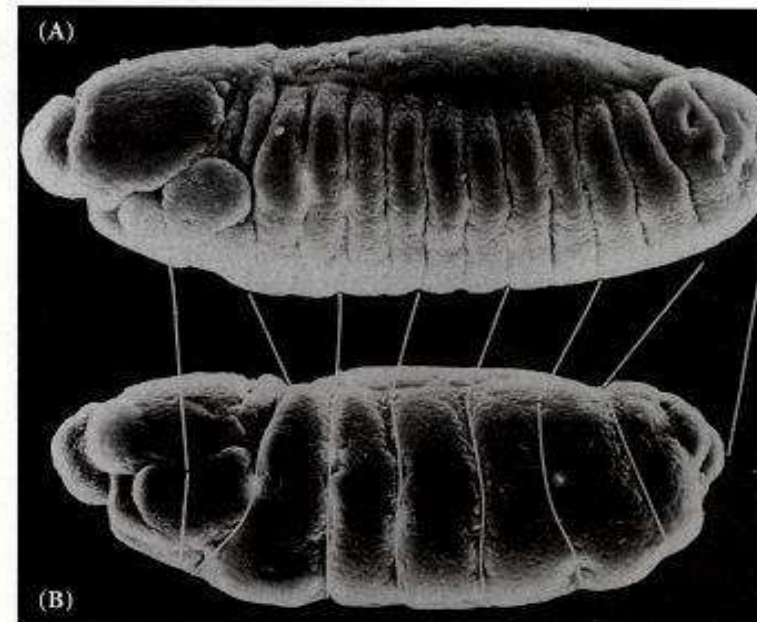
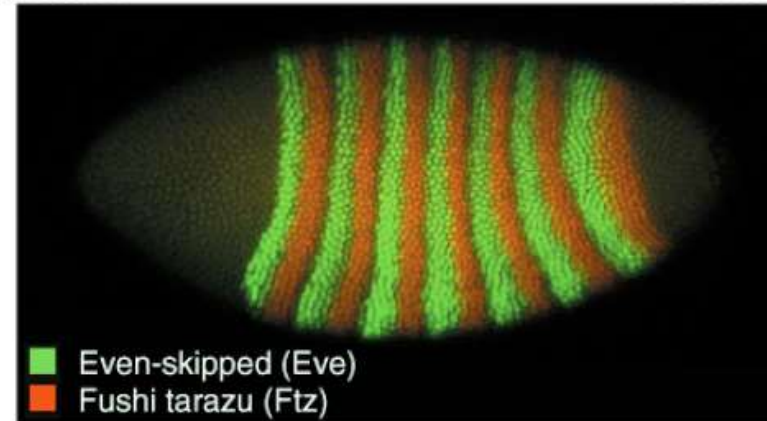
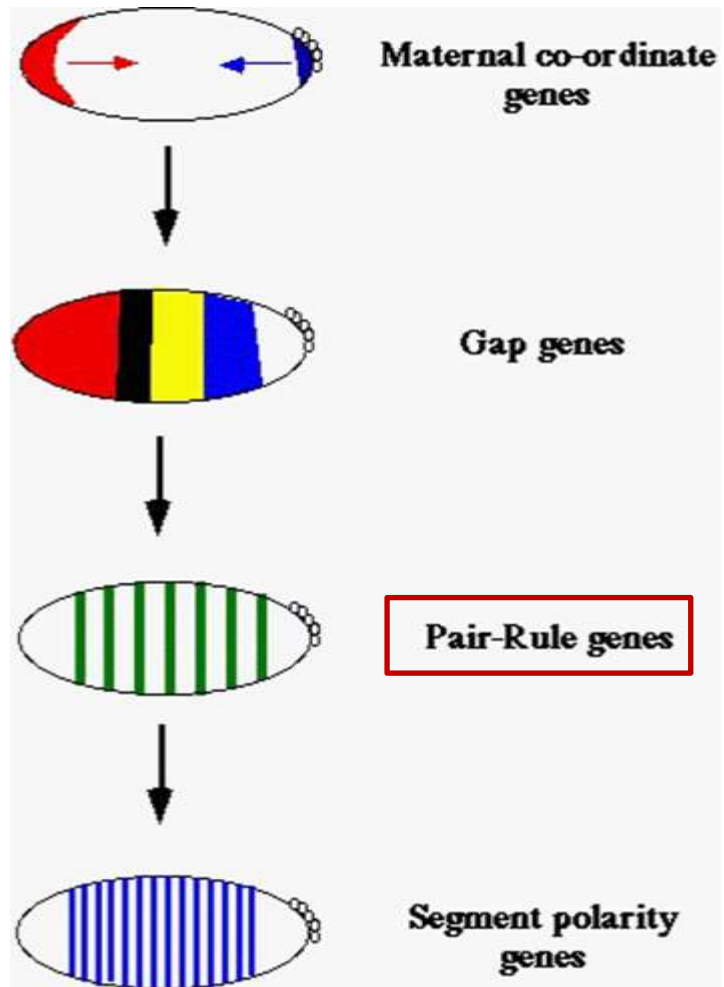
Gap genes

Zones of expression of four gap genes: *hunchback*, *Kruppel*, *knirps*, and *giant* in late syncytial blastoderm embryos

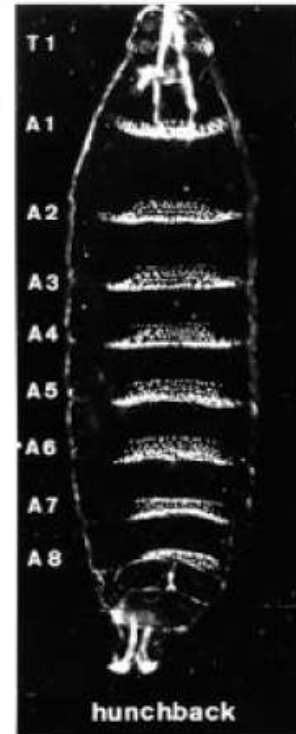
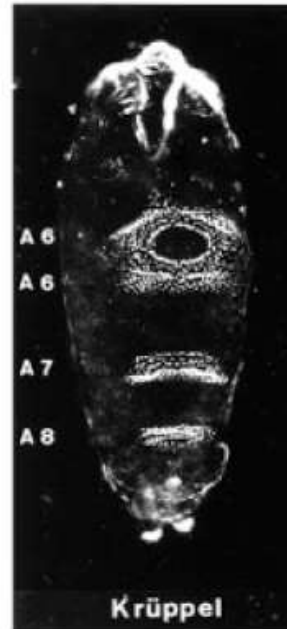
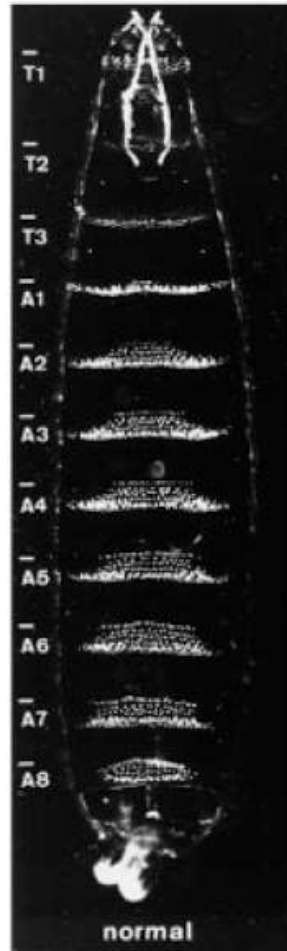
(a) Zones of gap gene expression



Pair-rule Genes



Defects in segmentation from mutations in gap genes



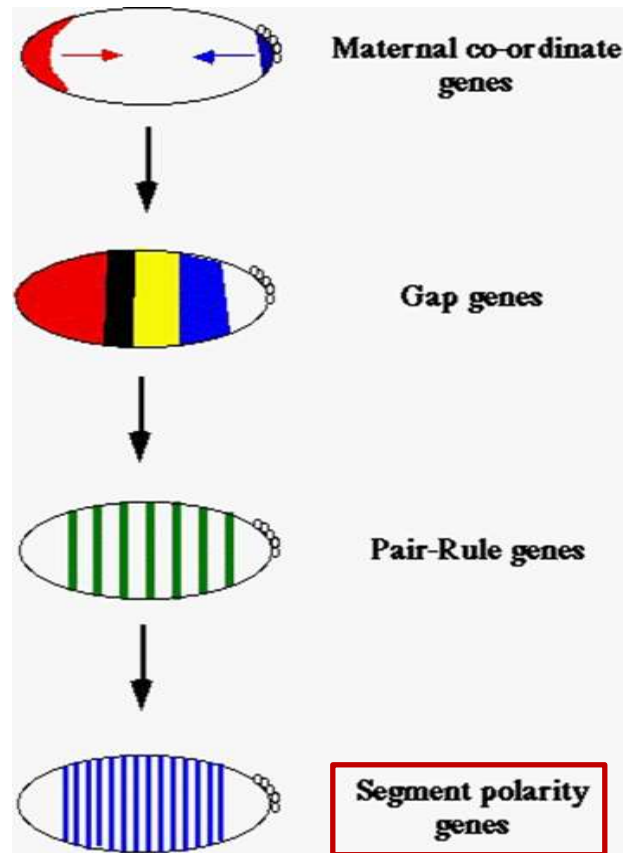
B)

C)

D.)

A)

Segment polarity genes are lowest level of segmentation hierarchy



Segment polarity genes

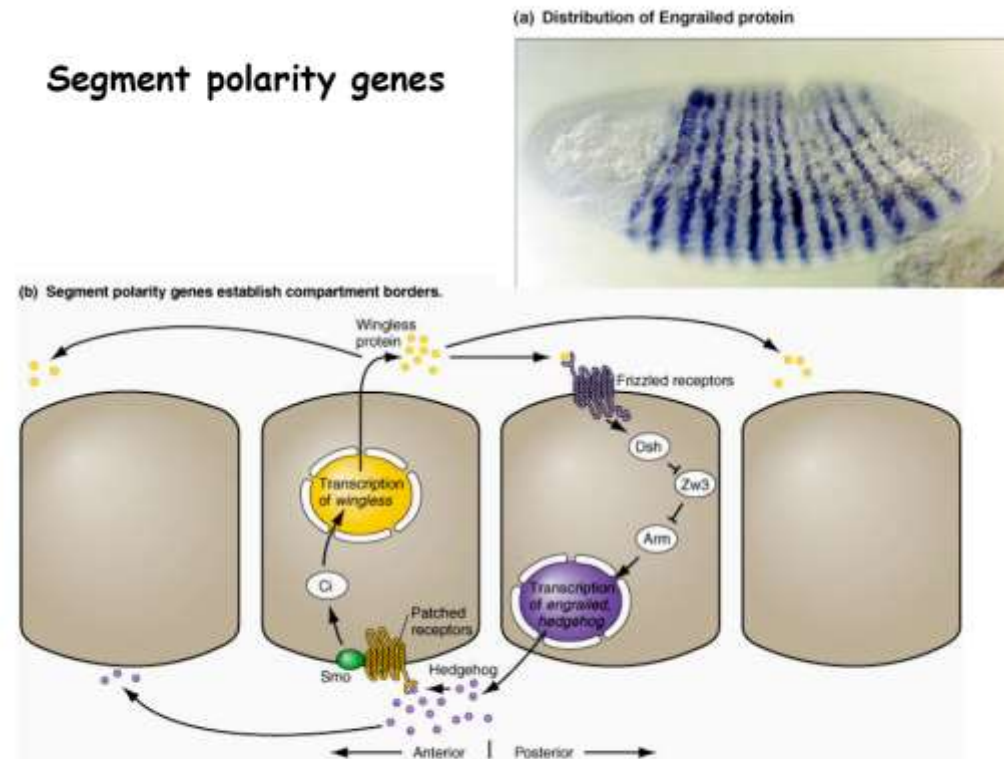
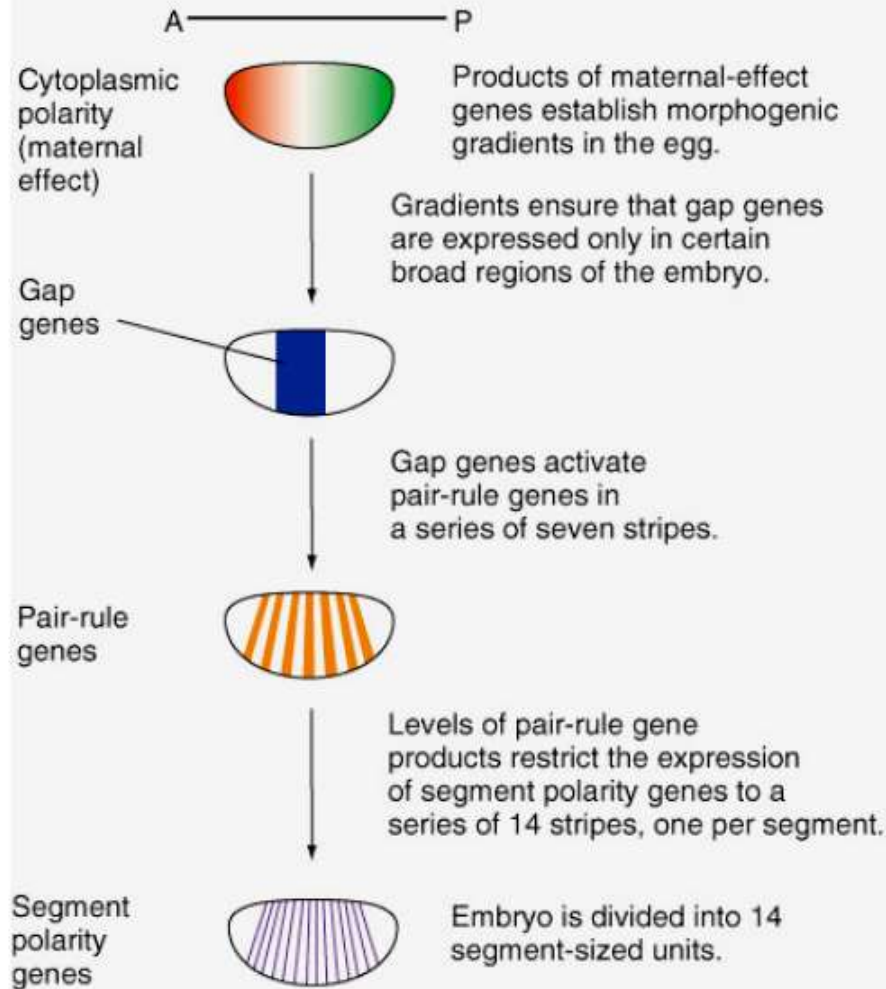


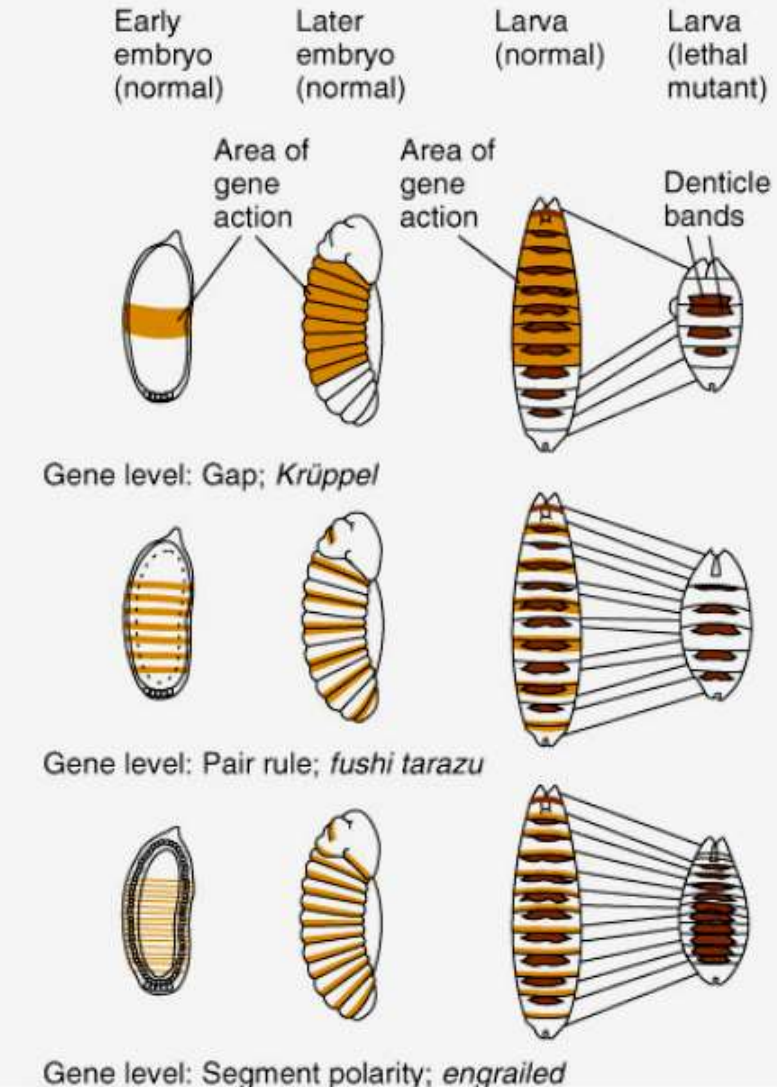
Fig. D.24

Drosophila embryogenesis: Segmentation genes

(a) The segmentation hierarchy



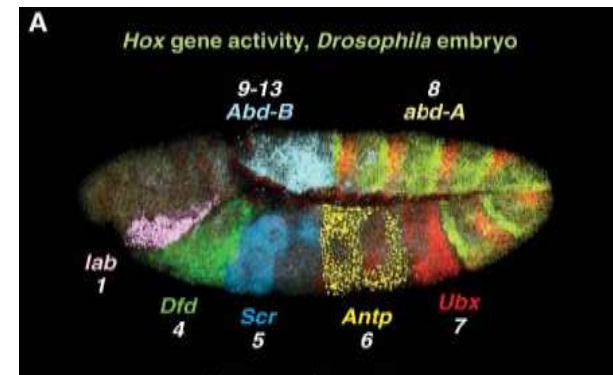
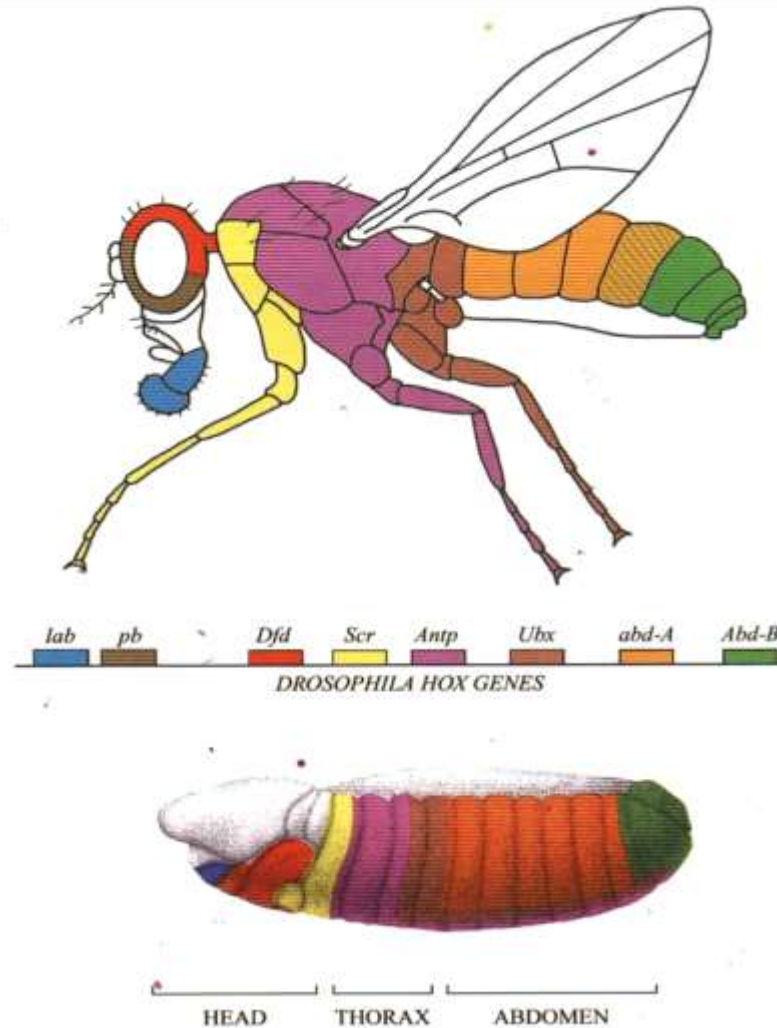
(b) Mutations in segmentation genes cause segment loss.



Homeotic genes: determine the fate of each segment



Edward B. Lewis



Drosophila Homeotic mutants

同源异形



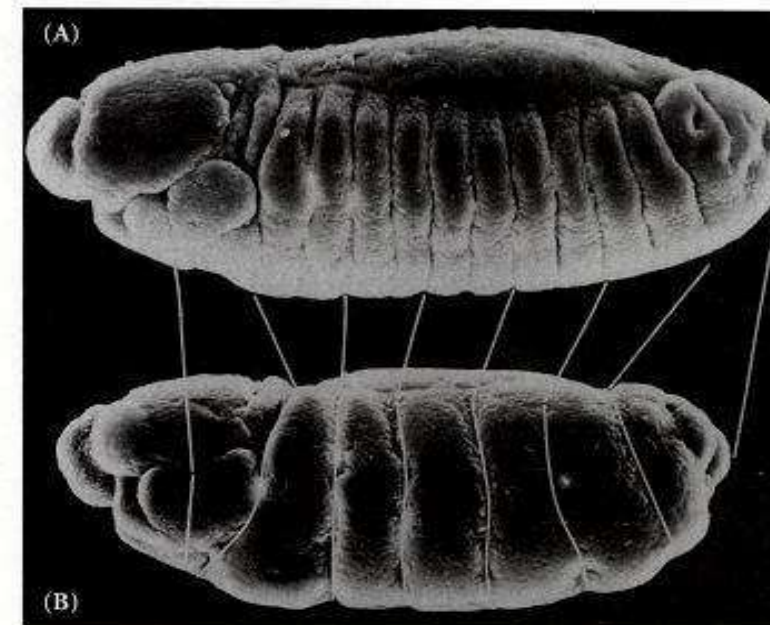
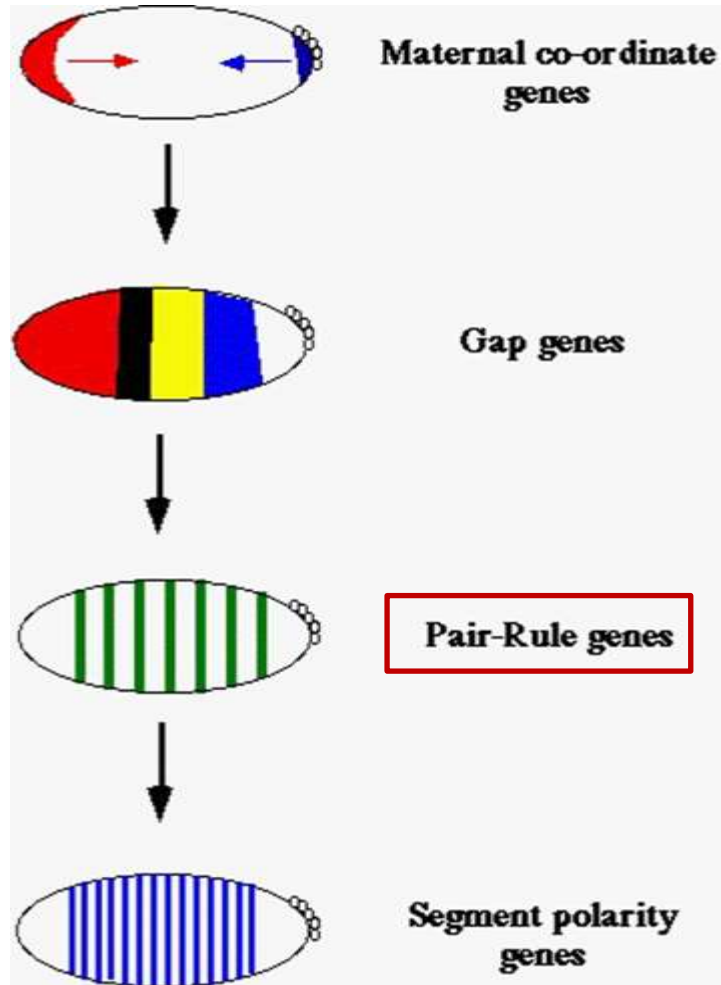
Edward B. Lewis

Wild type



Ubx

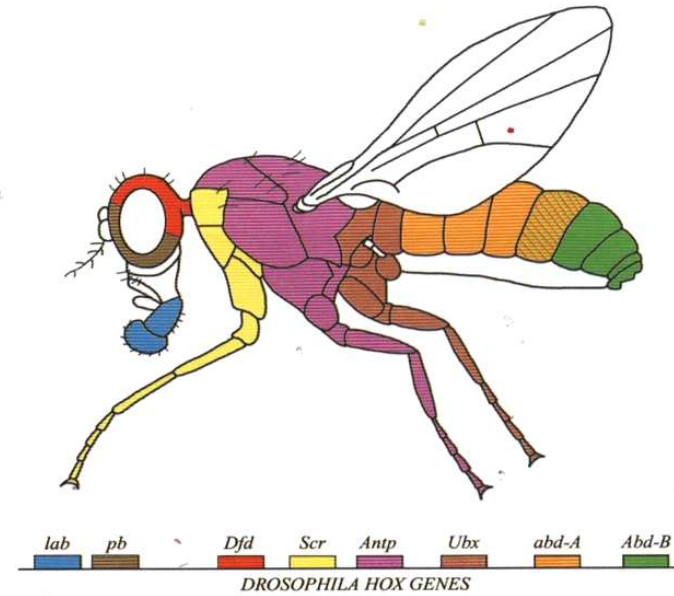
Pair-rule Genes



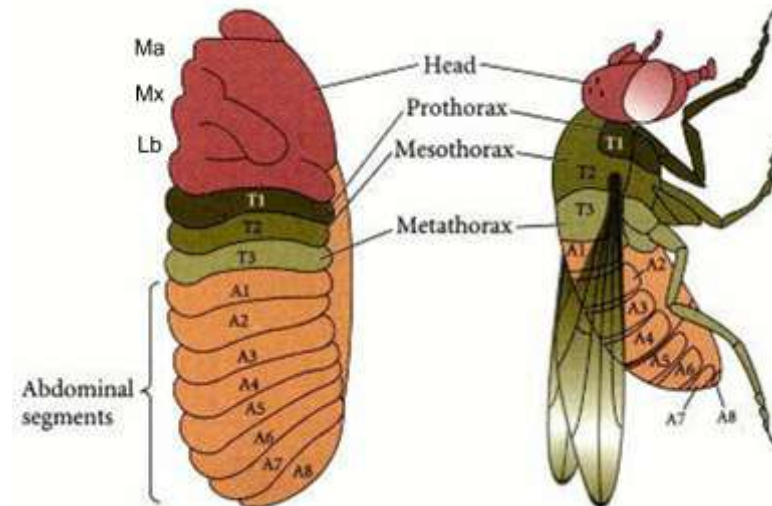
Drosophila Homeotic mutants

同源异形

Wild type

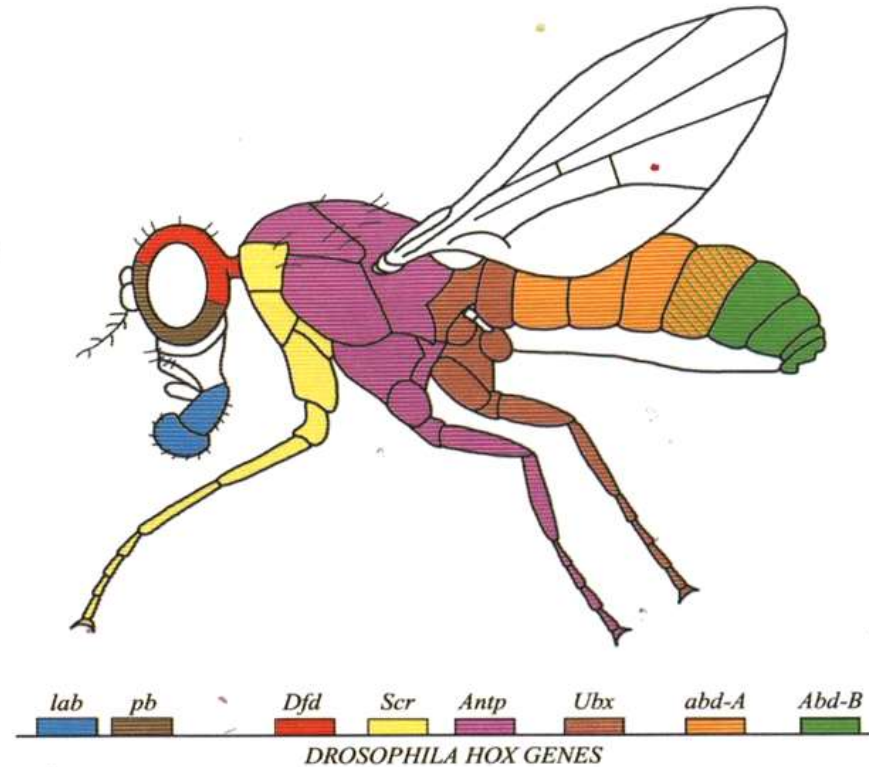
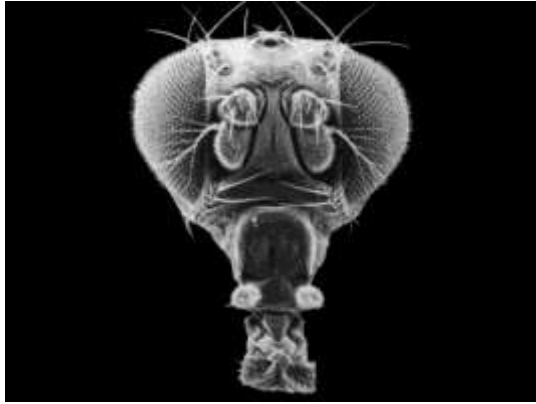


Ubx



Drosophila Homeotic mutants

同源异形



Antp: gain-of-function mutant

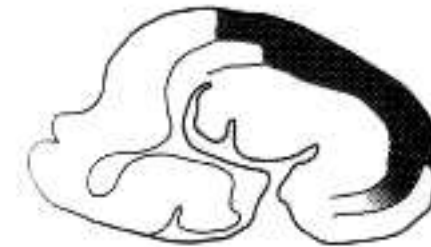
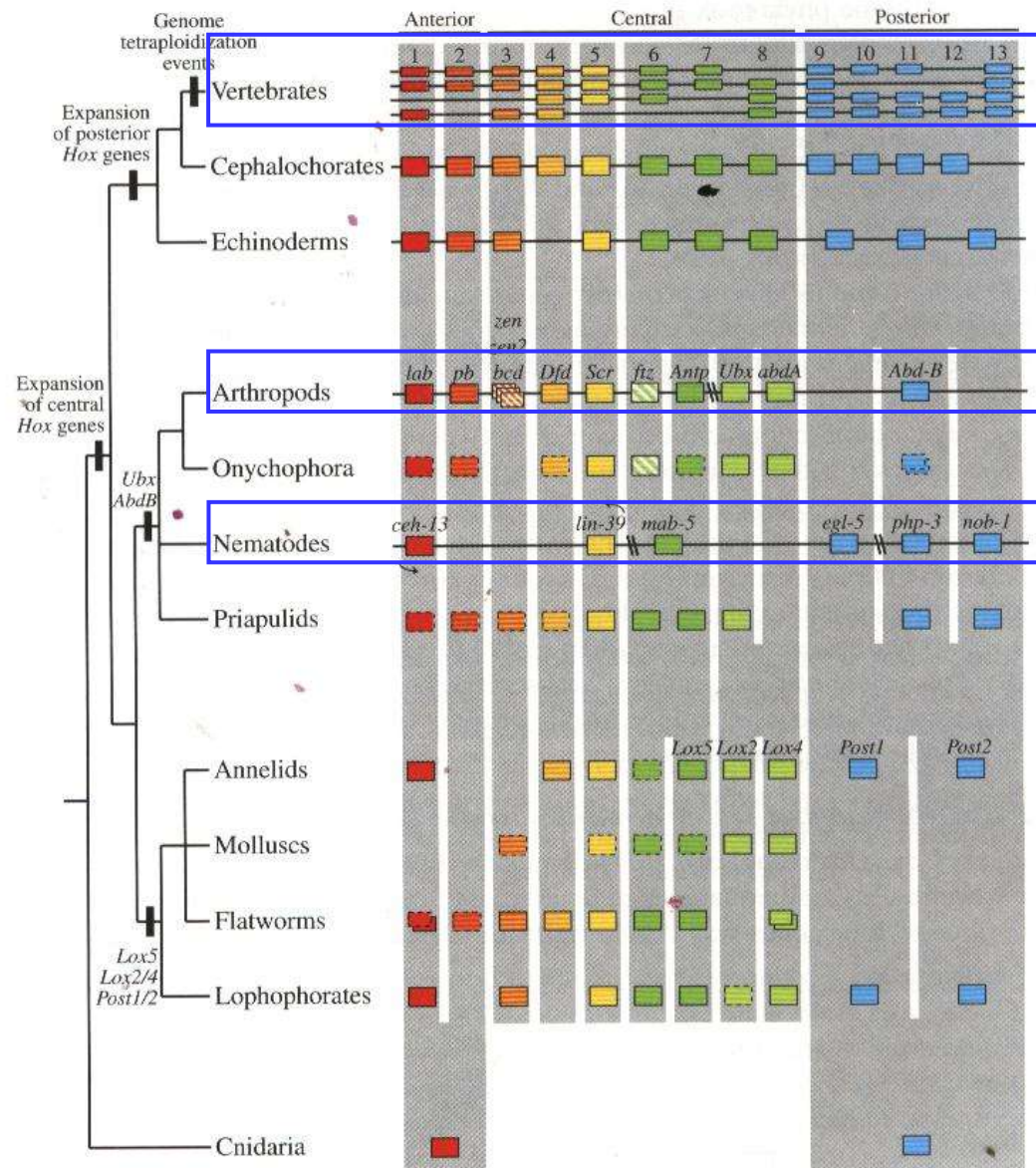
The homeotic genes contain a highly conserved homeobox (Hox)

lab	NNSGRTNFTNKQLTELEKEFHFNRYLTRARRRIEIANLTQLNETQVKIWFQNRRMKQKKRV
pb	PRRLRTAYTWTQLLELEKEFHFNRYLTRARRRIEIAASLTQLTERQVKIWFQNRRMKHKRQT
Dfd	PKRQRTAYTRHQLTELEKEFHFNRYLTRRRRIEIAHTLVLSERQIKIWFQNRRMKWKKN
Scr	TKRQRTSYTRYQTLELEKEFHFNRYLTRRRRIEIAHALCLTERQIKIWFQNRRMKWKKEH
Antp	RKRGRQTYTRYQTLELEKEFHFNRYLTRRRRIEIAHALCLTERQIKIWFQNRRMKWKKEN
Ubx	RRRGRQTYTRYQTLELEKEFHFNHYLTRRRRIEIAHALCLTERQIKIWFQNRRMKLKKEI
abd-A	RRRGRQTYTRYQTLELEKEFHFNHYLTRRRRIEIAHALCLTERQIKIWFQNRRMKLKKEI
abd-B	VRKKRKFPYSKFTLELEKEFHFNAYVSKQKRWELARNLTQLTERQVKIWFQNRRMKKNKNS
consensus	-RRGRT-YTR-QTLELEKEFHFNRYLTRRRRIEIAHALCLTERQIKIWFQNRRMK-KKE-
	Helix 1 Helix 2 Helix 3

The Homeobox (*Hox*) genes are transcription regulators

the Homeobox encodes a Homeodomain

Evolution of Metazoan *Hox* genes



mouse *Ubx*



Drosophila *Ubx*

刺胞动物: 珊瑚虫、水螅



The Nobel Prize in Physiology or Medicine 2011

"for their discoveries concerning the activation of innate immunity"



Jules Hoffmann

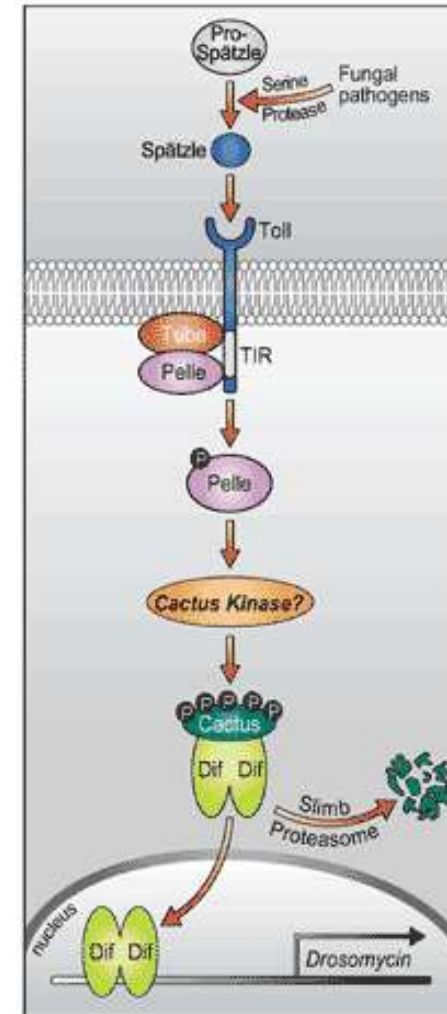
Toll/NF- κ B 信号通路

调控先天免疫

Cell, Vol. 86, 973–983, September 20, 1996, Copyright ©1996 by Cell Press

The Dorsoventral Regulatory Gene Cassette *spätzle/Toll/cactus* Controls the Potent Antifungal Response in *Drosophila* Adults

Bruno Lemaitre, Emmanuelle Nicolas, Lydia Michaut,
Jean-Marc Reichhart, and Jules A. Hoffmann





The Nobel Prize in Physiology or Medicine 2017

"for their discoveries of molecular mechanisms controlling the circadian rhythm"



Jeffrey C. Hall



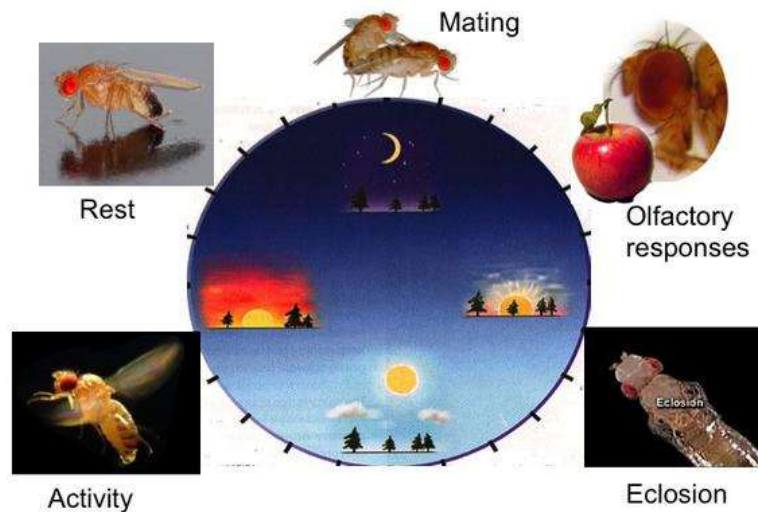
Michael Rosbash



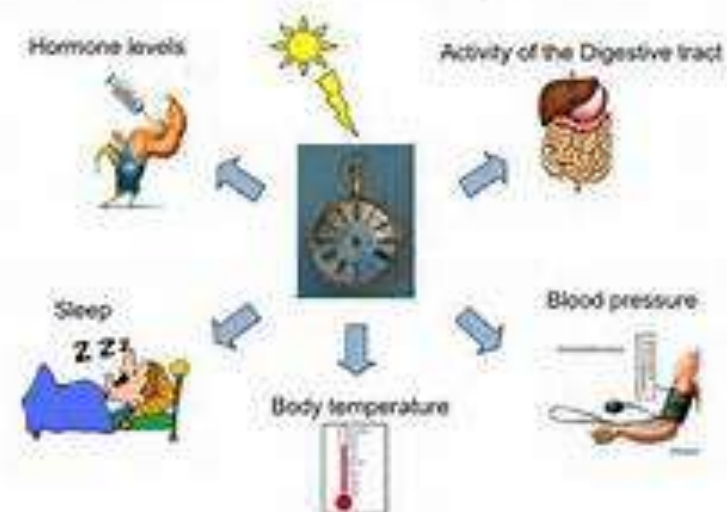
Michael W. Young



Drosophila Circadian rhythms



Human Circadian Rhythms





Seymour Benzer
1921 - 2007



Ronald Konopka
1947 - 2015

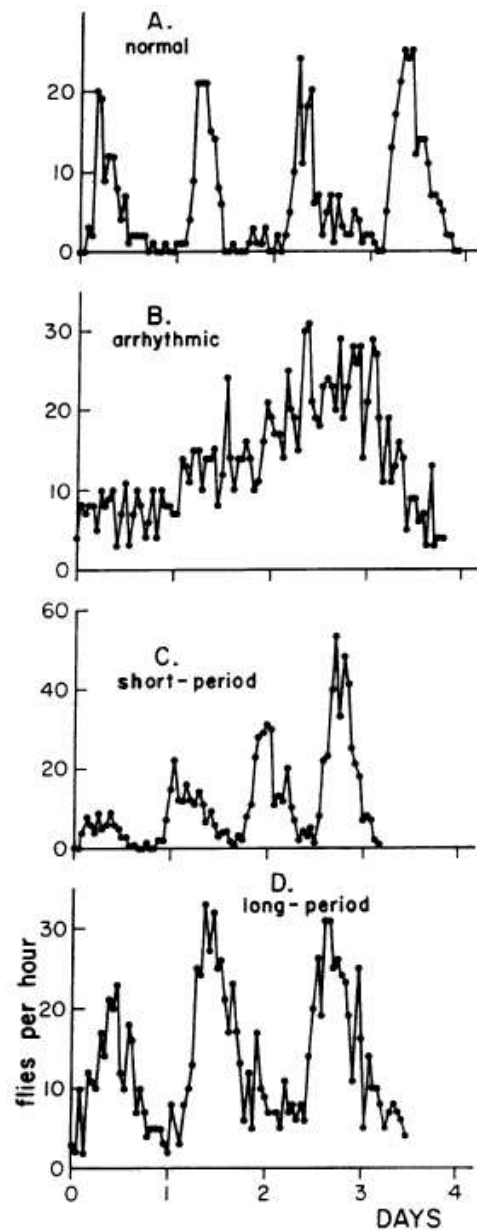
Proc. Nat. Acad. Sci. USA
Vol. 68, No. 9, pp. 2112-2116, September 1971

Clock Mutants of *Drosophila melanogaster* (eclosion/circadian/rhythms/X chromosome)

RONALD J. KONOPKA AND SEYMOUR BENZER

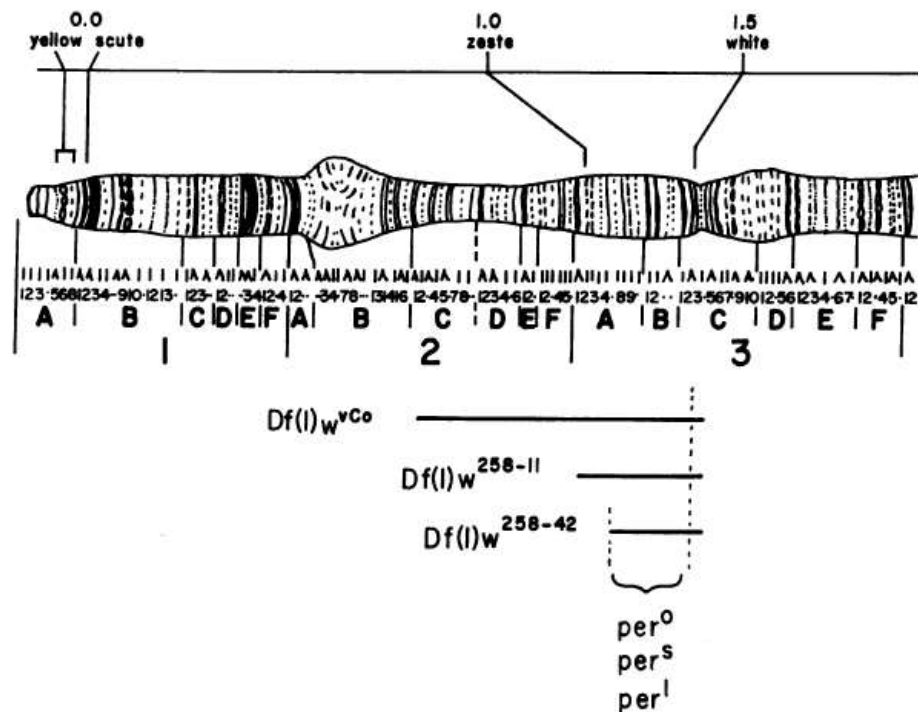
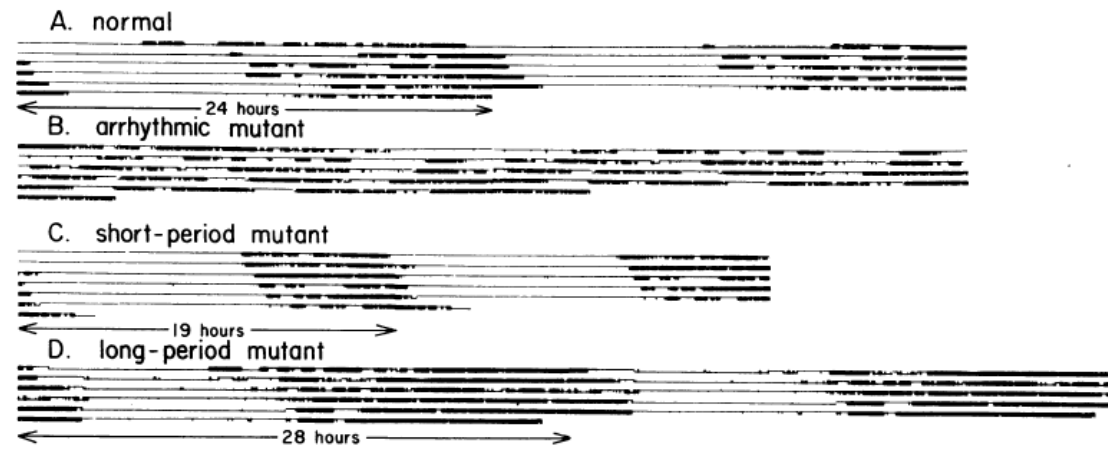
Division of Biology, California Institute of Technology, Pasadena, Calif. 91109

ABSTRACT Three mutants have been isolated in which the normal 24-hour rhythm is drastically changed. One mutant is arrhythmic; another has a period of 19 hr; a third has a period of 28 hr. Both the eclosion rhythm of a population and the locomotor activity of individual flies are affected. All these mutations appear to involve the same functional gene on the X chromosome.



Eclosion rhythms, in constant darkness,

Locomotor activity rhythms,



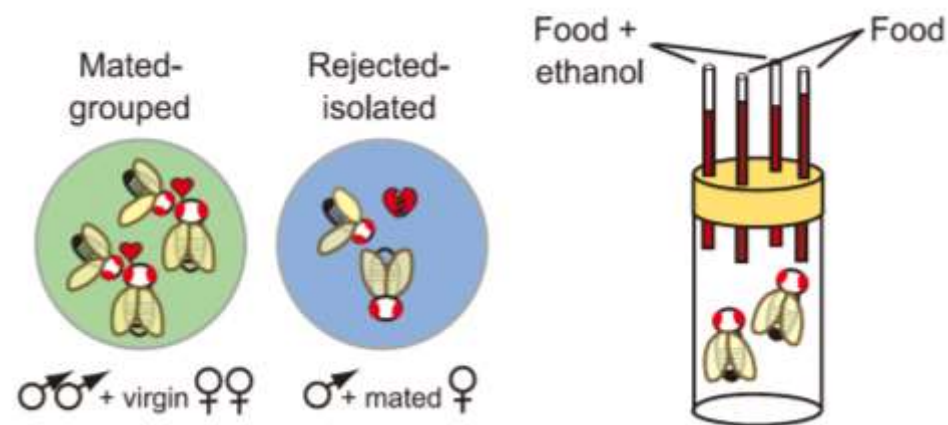
果蝇失恋也会“借酒消愁”

REPORT

Sexual Deprivation Increases Ethanol Intake in *Drosophila*

G. Shohat-Ophir^{*,†}, K. R. Kaun[†], R. Azanchi[†], H. Mohammed, U. Heberlein^{*,†}

Science 16 Mar 2012:



果蝇也“争强好胜”

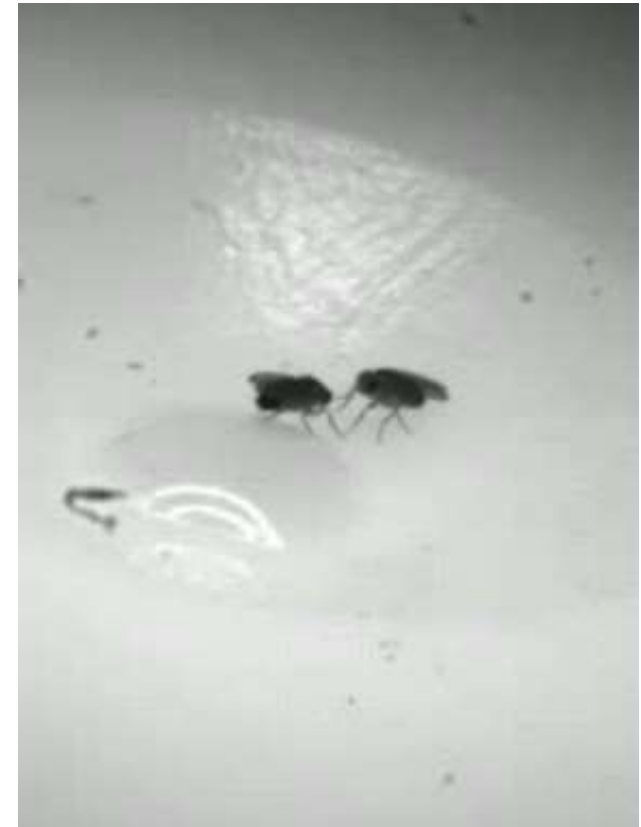
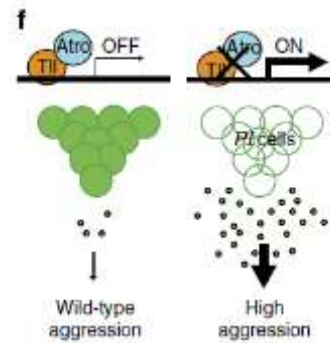
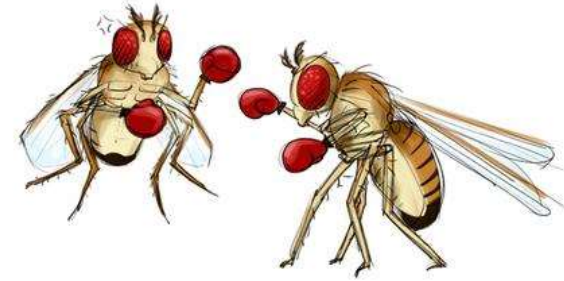
ARTICLE

Received 18 Jul 2013 | Accepted 23 Dec 2013 | Published 4 Feb 2014

DOI: 10.1038/ncomms4177

Tailless and Atrophin control *Drosophila* aggression by regulating neuropeptide signalling in the *pars intercerebralis*

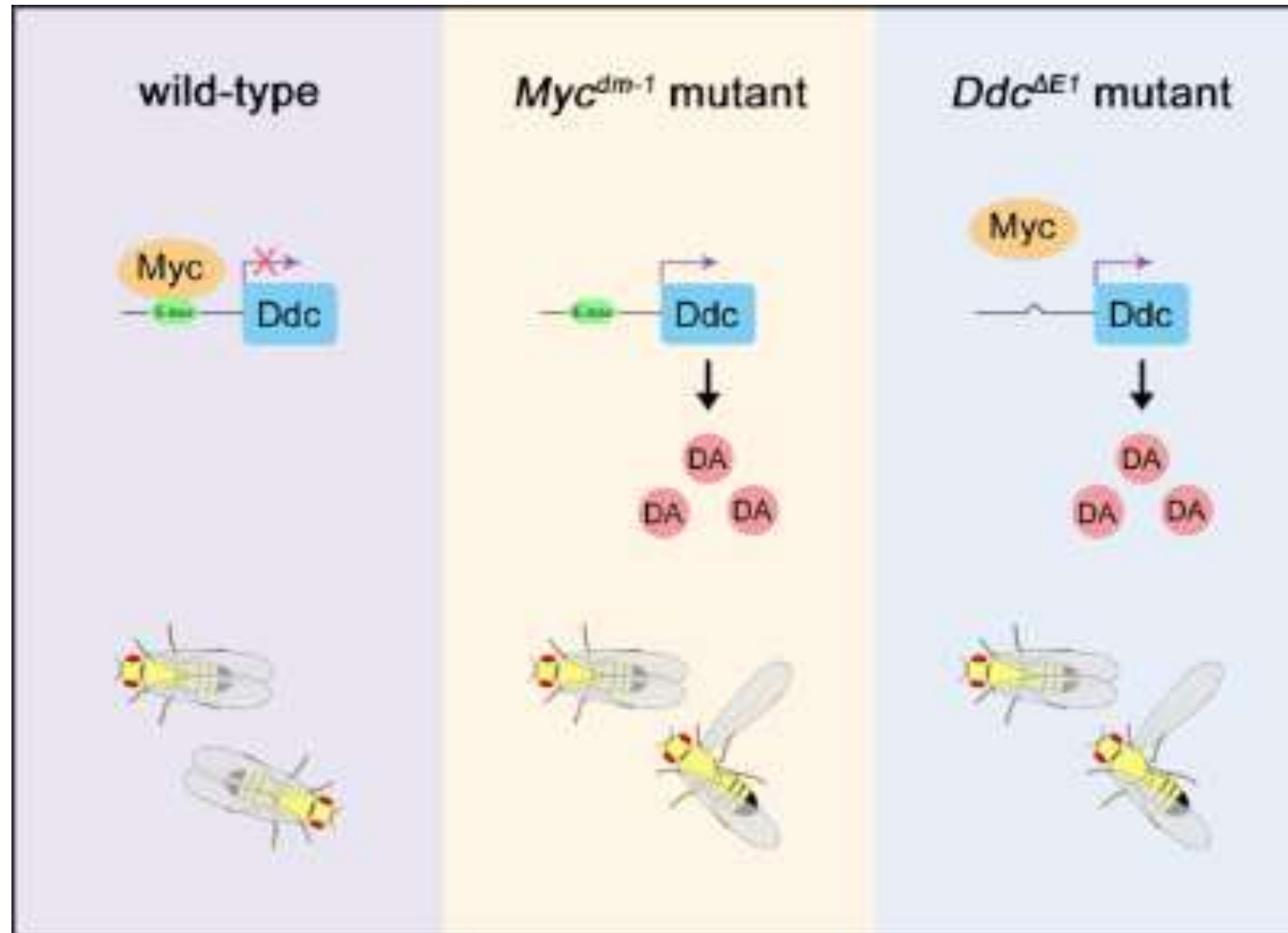
Shaun M. Davis^{1,*}, Amanda L. Thomas^{1,*}, Krystle J. Nomie^{1,*}, Longwen Huang³ & Herman A. Dierick^{1,2,3,4}





Myc suppresses male-male courtship in *Drosophila*

Yu Pan¹, Wanzhen Li¹, Zhu Deng², Yihao Sun³, Xianjue Ma⁴, Ruijuan Liang¹, Xiaowei Guo¹, Ying Sun¹, Wenzhe Li¹, Renjie Jiao², Lei Xue^{1 3}



EMBO J：薛雷课题组揭示果蝇同性求偶行为的新机制

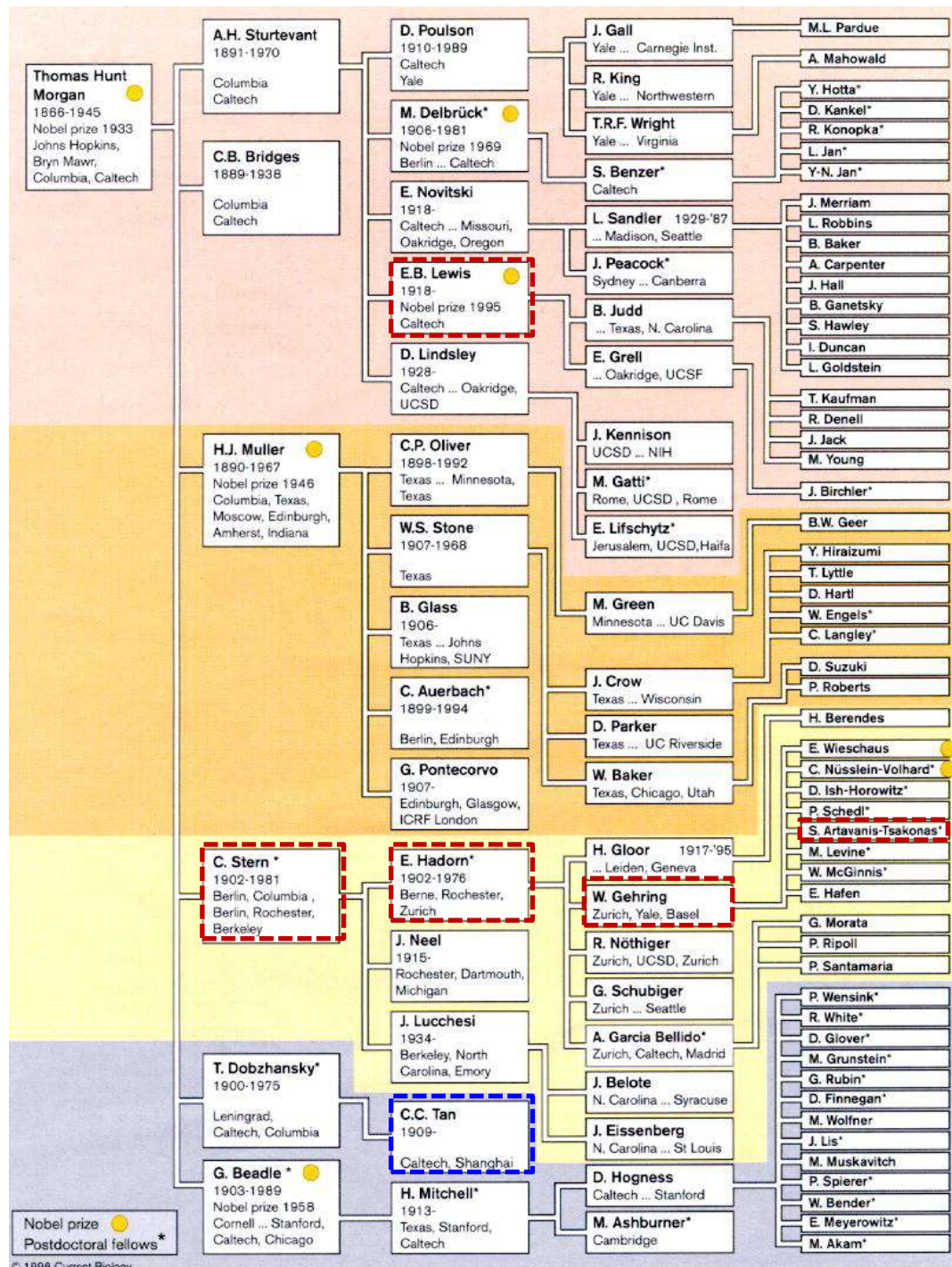
2022年02月28日 07:57 新浪网 作者 brainnews脑科学新闻

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薛雷课题组揭示果蝇同性求偶行为的新机制

2022-02-27 18:23:58 来源: BioArt





Homeotic mutants

The Morgan pedigree

Segmentation genes: maternal, gap, pair-rule, segment polarity

Homeobox