

The place of *Drosophila* among the metazoans

2

Peter Holland
Zoology, Oxford

24 March 2000



24 March 2000

The Genome Sequence of *Drosophila melanogaster*

Mark D. Adams,^{1*} Susan E. Celniker,² Robert A. Holt,¹ Cheryl A. Evans,¹ Jeannine D. Gocayne,¹ Peter G. Amanatides,¹ Steven E. Scherer,³ Peter W. Li,¹ Roger A. Hoskins,² Richard F. Galle,² Reed A. George,² Suzanna E. Lewis,⁴ Stephen Richards,² Michael Ashburner,⁵ Scott N. Henderson,¹ Granger G. Sutton,¹ Jennifer R. Wortman,¹ Mark D. Yandell,¹ Qing Zhang,¹ Lin X. Chen,¹ Rhonda C. Brandon,¹ Yu-Hui C. Rogers,¹ Robert G. Blazej,² Mark Champe,² Barret D. Pfeiffer,² Kenneth H. Wan,² Clare Doyle,² Evan G. Baxter,² Gregg Helt,⁶ Catherine R. Nelson,⁴ George L. Gabor Miklos,⁷ Josep F. Abril,⁸ Anna Agbayani,² Hui-Jin An,¹ Cynthia Andrews-Pfannkoch,¹ Danita Baldwin,¹ Richard M. Ballew,¹ Anand Basu,¹ James Baxendale,¹ Leyla Bayraktaroglu,⁹ Ellen M. Beasley,¹ Karen Y. Beeson,¹ P. V. Benos,¹⁰ Benjamin P. Berman,² Deepali Bhandari,¹ Slava Bolshakov,¹¹ Dana Borkova,¹² Michael R. Botchan,¹³ John Bouck,³ Peter Brokstein,⁴ Phillipe Brottier,¹⁴ Kenneth C. Burtis,¹⁵ Dana A. Busam,¹ Heather Butler,¹⁶ Edouard Cadieu,¹⁷ Angela Center,¹ Ishwar Chandra,¹ J. Michael Cherry,¹⁸ Simon Cawley,¹⁰ Carl Dahlke,¹ Lionel B. Davenport,¹ Peter Davies,¹ Beatriz de Pablos,²⁰ Arthur Delcher,¹ Zuoming Deng,¹ Anne Deslattes Mays,¹ Ian Dew,¹ Suzanne M. Dietz,¹ Kristina Dodson,¹ Lisa E. Doup,¹ Michael Downes,²¹ Shannon Dugan-Rocha,³ Boris C. Dunkov,²² Patrick Dunn,¹ Kenneth J. Durbin,³ Carlos C. Evangelista,¹ Concepcion Ferraz,²³ Steven Ferreira,¹ Wolfgang Fleischmann,⁵ Carl Fosler,¹ Andrei E. Gabrielian,¹ Neha S. Garg,¹ William M. Gelbart,⁹ Ken Glasser,¹ Anna Glodek,¹ Fangcheng Gong,¹ J. Harley Gorrell,³ Zhiping Gu,¹ Ping Guan,¹ Michael Harris,¹ Nomi L. Harris,² Damon Harvey,⁴ Thomas J. Heiman,¹ Judith R. Hernandez,³ Jarrett Houck,¹ Damon Hostin,¹ Kathryn A. Houston,² Timothy J. Howland,¹ Ming-Hui Wei,¹ Chinyere Ibegwam,¹ Mena Jalali,¹ Francis Kalush,¹ Gary H. Karpen,²¹ Zhaoxi Ke,¹ James A. Kennison,²⁴ Karen A. Ketchum,¹ Bruce E. Kimmel,² Chinnappa D. Kodira,¹ Cheryl Kraft,¹ Saul Kravitz,¹ David Kulp,⁶ Zhongwu Lai,¹ Paul Lasko,²⁵ Yiding Lei,¹ Alexander A. Levitsky,¹ Jiayin Li,¹ Zhenya Li,¹ Yong Liang,¹ Xiaoying Lin,²⁶ Xiangjun Liu,¹ Bettina Mattei,¹ Tina C. McIntosh,¹ Michael P. McLeod,³ Duncan McPherson,¹ Gennady Merkulov,¹ Natalia V. Milshina,¹ Clark Mobarry,¹ Joe Morris,⁶ Ali Moshrefi,² Stephen M. Mount,²⁷ Mee Moy,¹ Brian Murphy,¹ Lee Murphy,²⁸ Donna M. Muzny,³ David L. Nelson,³ David R. Nelson,²⁹ Keith A. Nelson,¹ Katherine Nixon,² Deborah R. Nusskern,¹ Joanne M. Paclab,² Michael Palazzolo,² Gjange S. Pittman,¹ Sue Pan,¹ John Pollard,¹ Vinita Puri,¹ Martin G. Reese,⁴ Knut Reinert,¹ Karin Remington,¹ Robert D. C. Saunders,³⁰ Frederick Scheeler,¹ Hua Shen,³ Bixiang Christopher Shue,¹ Inga Sidén-Kiamos,¹¹ Michael Simpson,¹ Marian P. Skupski,¹ Tom Smith,¹ Eugene Spier,¹ Allan C. Spradling,³¹ Mark Stapleton,² Renee Strong,¹ Eric Sun,¹ Robert Svirskas,³² Cyndee Tector,¹ Russell Turner,¹ Eli Venter,¹ Aihui H. Wang,¹ Xin Wang,¹ Zhen-Yuan Wang,¹ David A. Wassarman,³³ George M. Weinstock,³ Jean Weissenbach,¹⁴ Sherita M. Williams,¹ Trevor Woodage,¹ Kim C. Worley,³ David Wu,¹ Song Yang,² Q. Alison Yao,¹ Jane Ye,¹ Ru-Fang Yeh,¹⁰ Jayshree S. Zaveri,¹ Ming Zhan,¹ Guangren Zhang,¹ Qi Zhao,¹ Liansheng Zheng,¹ Xiangqun H. Zheng,¹ Fei N. Zhong,¹ Wenyan Zhong,¹ Xiaojun Zhou,³ Shiaoqing Zhu,¹ Xiaohong Zhu,¹ Hamilton O. Smith,¹ Richard A. Gibbs,³ Eugene W. Myers,¹ Gerald M. Rubin,³⁴ J. Craig Venter¹

Science 287, 2185 - 2195

Metazoan genomes published (up to 2006)

Pictures from here removed for copyright reasons.

Human Chimp Mouse Rat Dog Chicken *Takifugu* *Tetraodon*

Anopheles *Apis*

Ciona intestinalis *Oikopleura dioica*

Drosophila melanogaster & *pseudoobscura*

Caenorhabditis elegans & *briggsae*

Other essentially 'complete' Metazoan genomes

Pictures from here removed for copyright reasons.

Macaque

Cow

Opposum

Xenopus

Zebrafish

Stickleback

Sea urchin

Schmidtea

Lottia

Amphioxus

Ciona savignyi

Medaka

Several nematodes,
notably *Caenorhabditis remanei* & *Brugia*

Nematostella

10 more *Drosophila* species

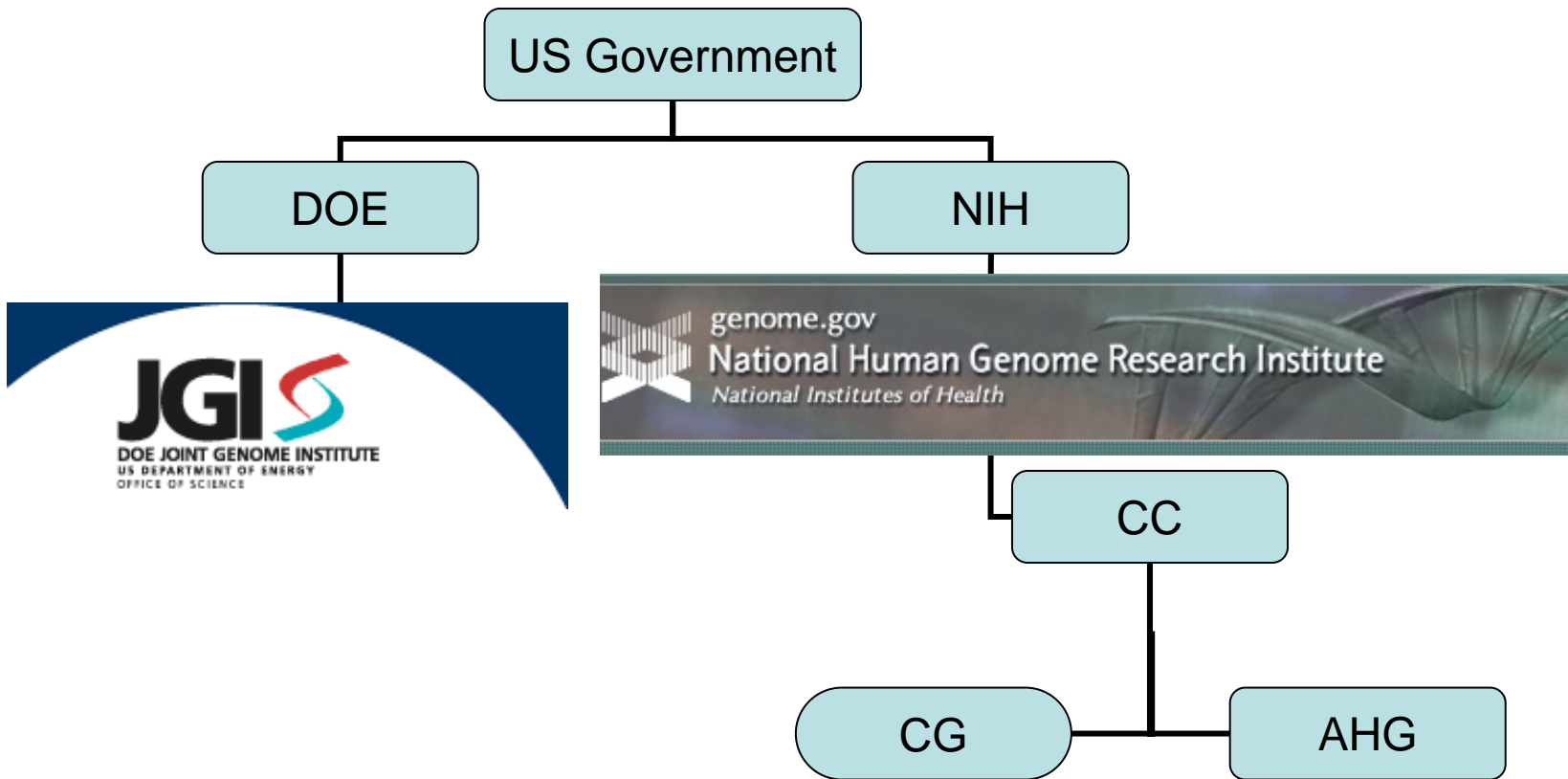
Daphnia

Tribolium

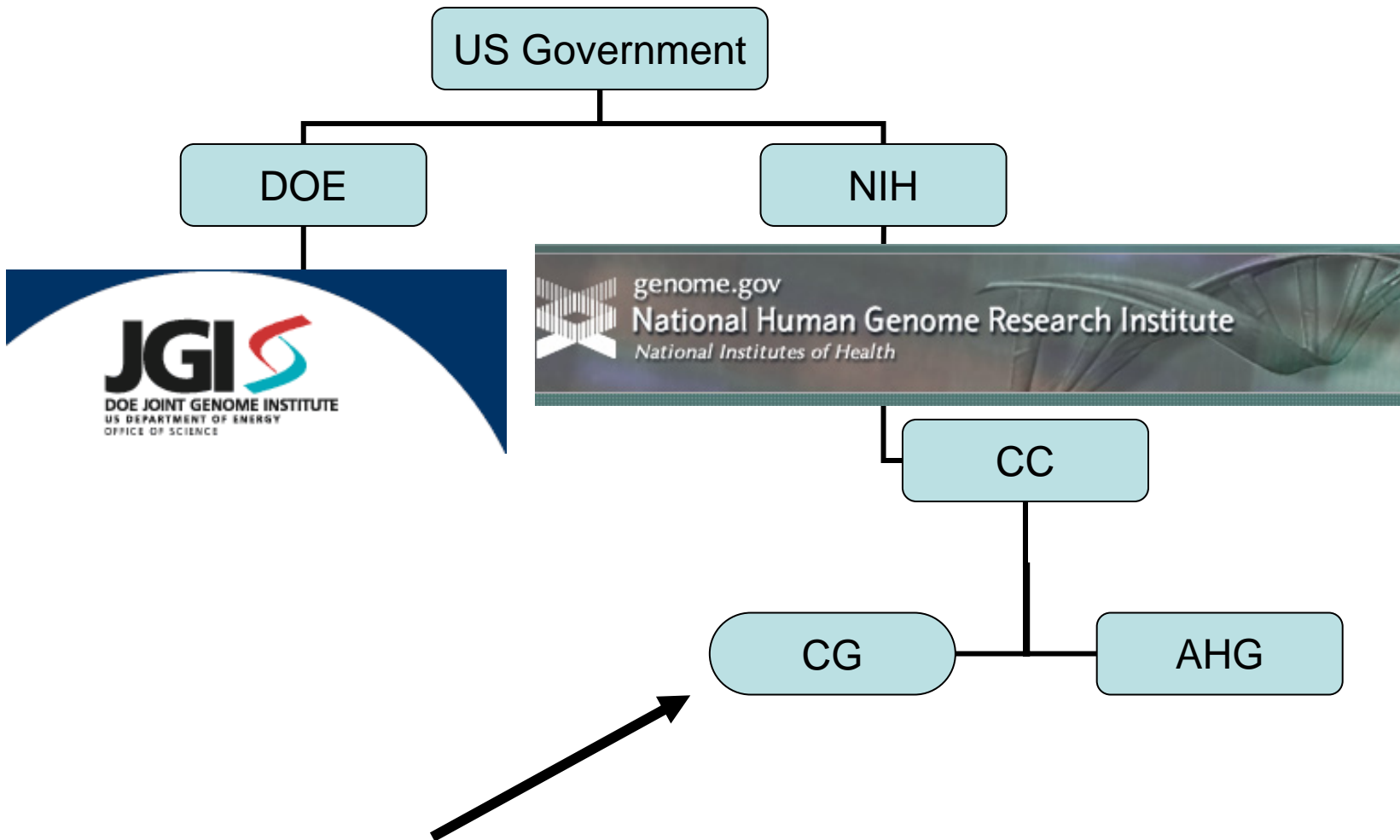
Nasonia

Sponge

Genome Sequencing



Genome Sequencing



Mammals – as of 2006

- **Placentals**

- Human, chimp, mouse, rat, dog = published
- Orang utan NHGRI 7x in progress
- Gibbon NHGRI, approved ←
- Macaque draft assembly released
- Marmaset NHGRI 2x approved, not started
- Bat NHGRI, 7x approved ←
- Cat NHGRI 2x in progress, upgraded to 7x plan ←
- Cow draft assembly released
- Elephant NHGRI 2x approved, upgraded to 7x plan ←
- Shrew NHGRI 2x approved, not started
- Hedgehog NHGRI 2x approved, not started
- Guinea pig NHGRI 2x approved, upgraded to 7x plan ←
- Tenrec NHGRI 2x approved, not started
- Armadillo NHGRI 2x approved, upgraded to 7x plan ←
- Rabbit NHGRI 2x approved, upgraded to 7x plan ←
- Tree shrew NHGRI, 7x approved ←

Summer 2006 decisions

- **Marsupials**

- Opossum draft assembly released
- Wallaby NHGRI 2x approved, not started

- **Monotremes**

- Platypus NHGRI draft assembly, started

Other vertebrates

Autumn 2006 decisions

- **Birds**

- Chicken – done 2005
- Zebra finch – now approved, underway 2006

- **Reptiles**

- Anolis lizard – NHGRI approved 2005
- Freshwater turtle – NHGRI approved 2006

- **Amphibians**

- *Xenopus tropicalis* done 2006

- **Basal sarcopterygians**

- *Coelacanth* approved 2006

- **Ray-finned fish**

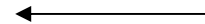
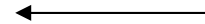
- Three-spined stickleback Almost done
- Takifugu Done
- Tetraodon Genoscope done 2005
- Zebrafish Sanger, almost done 2006
- Medaka almost done 2006
- Gar approved 2006

- **Cartilagenous fish**

- Skate *Raja erinacea* NHGRI approved
- Ratfish NHGRI provisionally approved

- **Jawless vertebrates**

- Sea lamprey Approved, problems
- Hagfish Approved 2006



Bilateria

Diploblasts



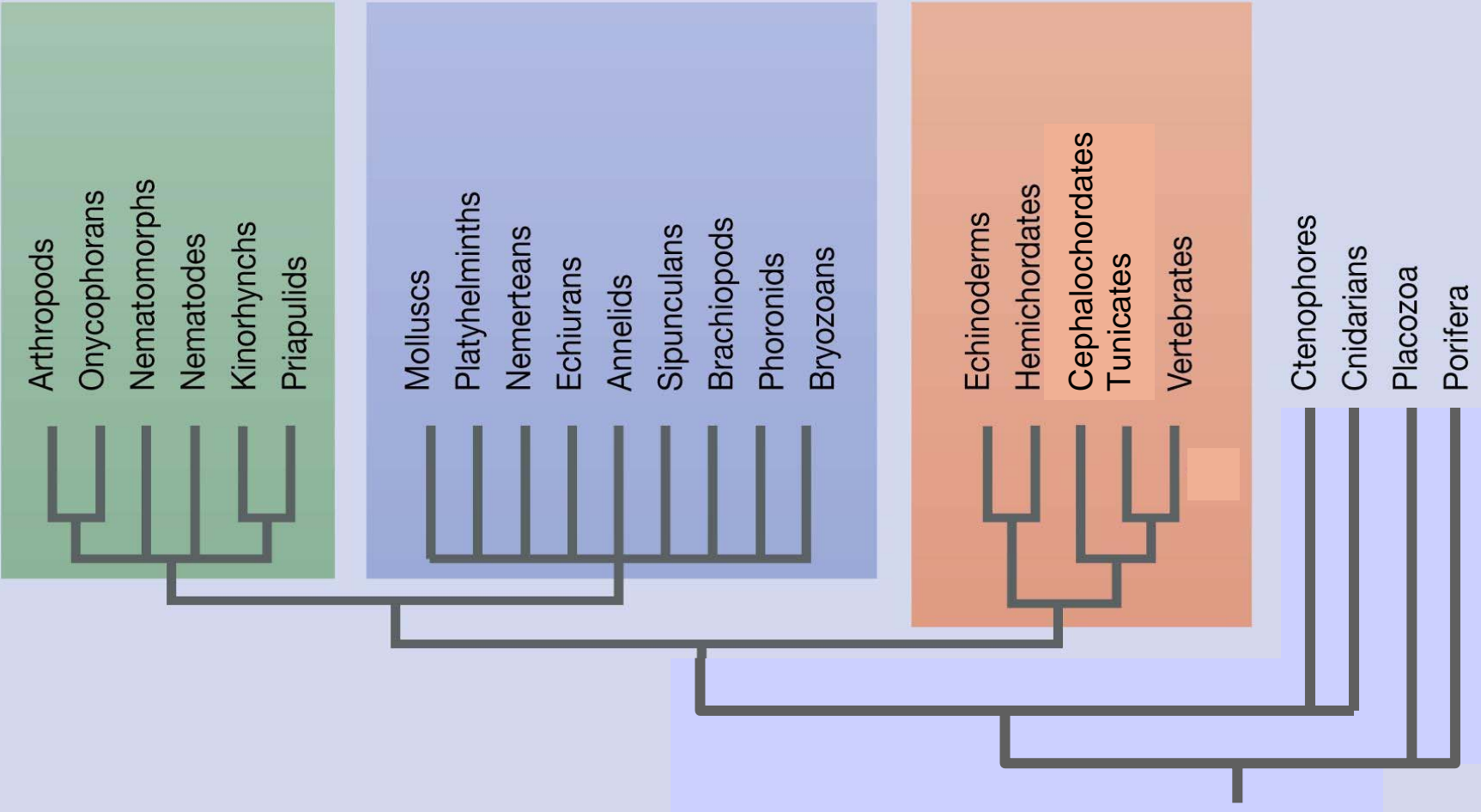
Ecdysozoans



Lophotrochozoans



Deuterostomes



Bilateria

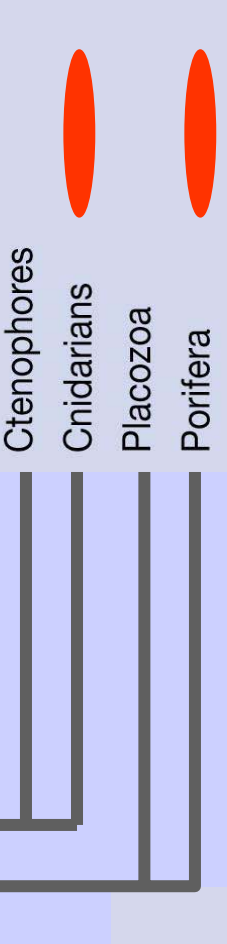
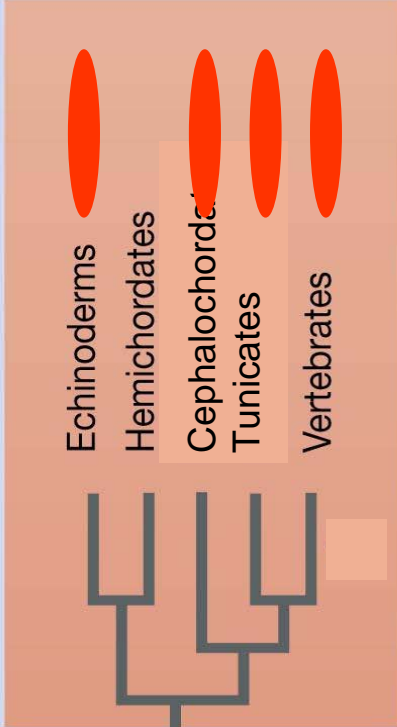
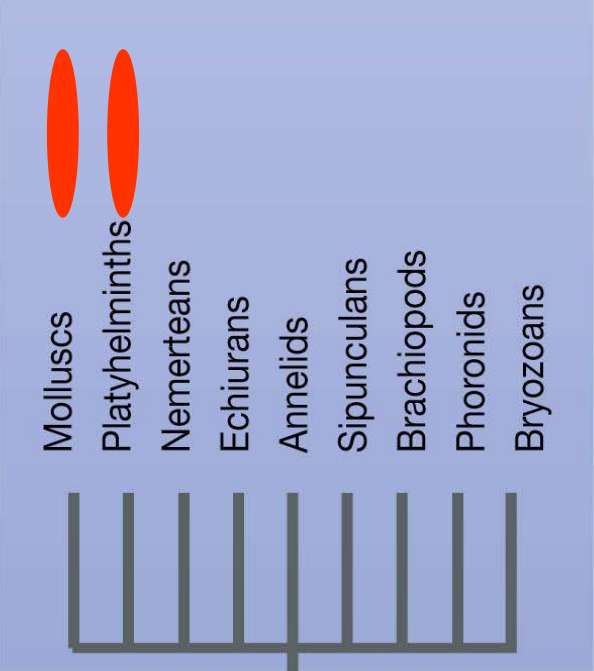
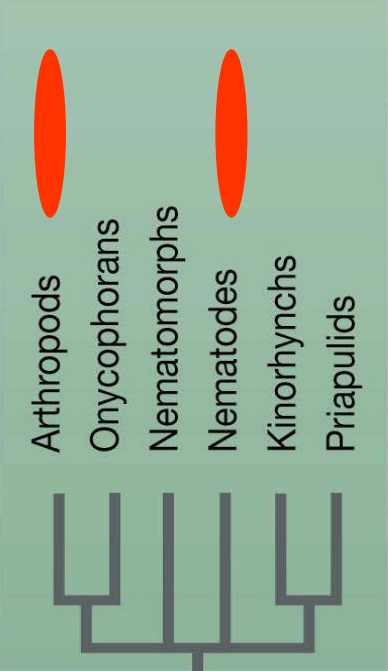
Diploblasts



Ecdysozoans

Lophotrochozoans

Deuterostomes



Phylum Arthropoda

Pictures from here removed for copyright reasons

e.g. spider

millipede

shrimp, Daphnia

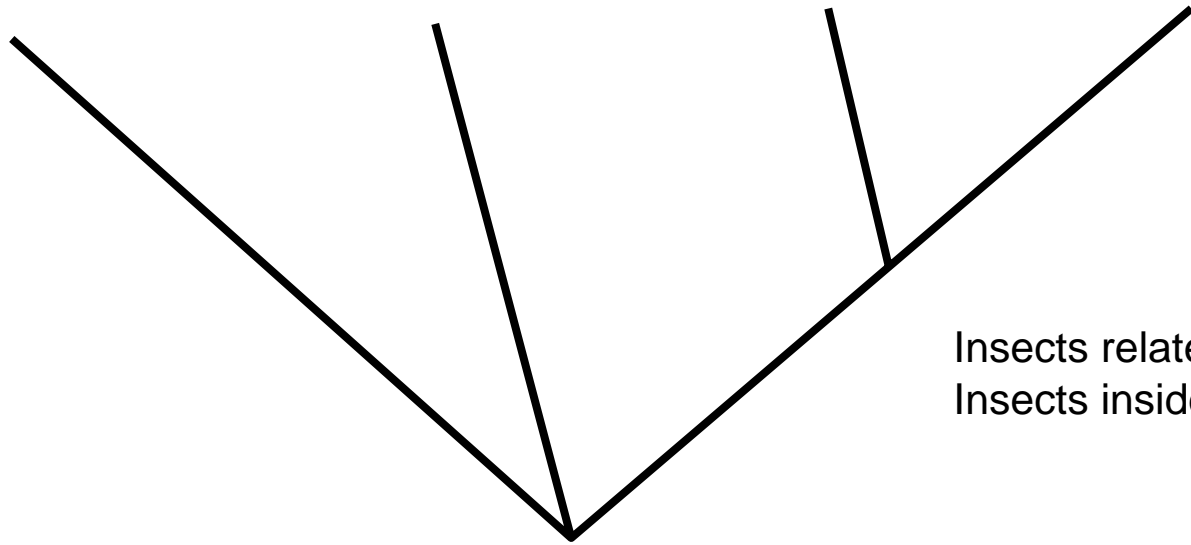
fly

Chelicerata

Myriapoda

Crustacea

Insecta



Insects related to Crustacea
Insects inside Crustacea?

Revised (molecular) phylogeny

Phylum Arthropoda

Pictures from here removed for copyright reasons

e.g. spider

millipede

Daphnia

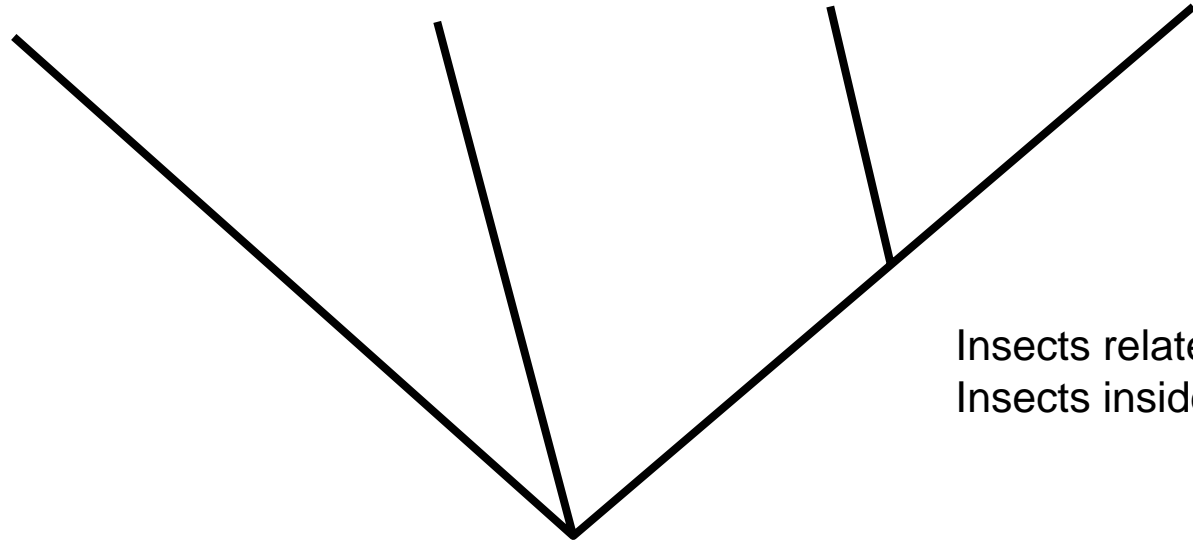
fly

Chelicerata

Myriapoda

Crustacea

Insecta



Insects related to Crustacea
Insects inside Crustacea?

Revised (molecular) phylogeny

Class Insecta

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e.g. bristletail

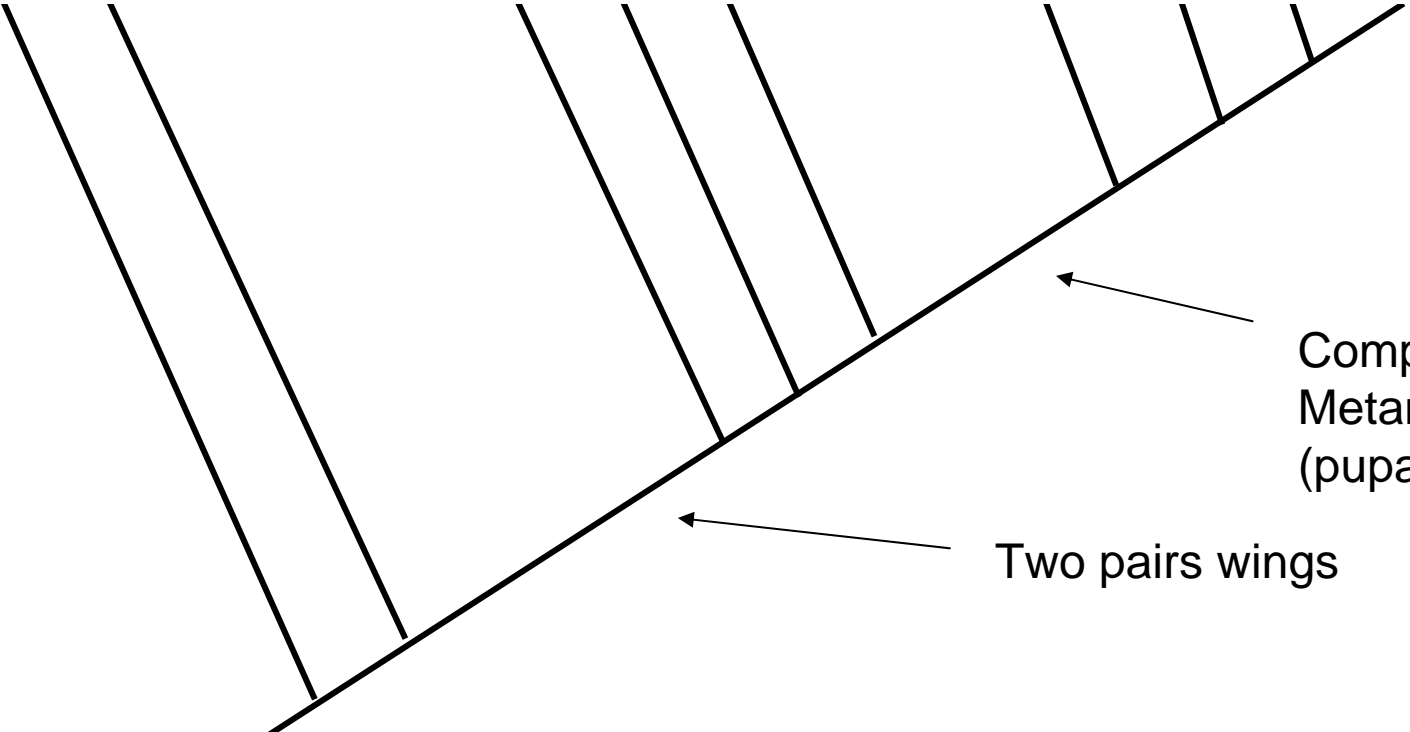
pea aphid

fly

Wingless insects

Hemimetabola

Holometabola



Complete Metamorphosis (pupa)

Two pairs wings

Class Insecta

Pea aphid – in progress

Pictures from here removed for copyright reasons

e.g. bristletail

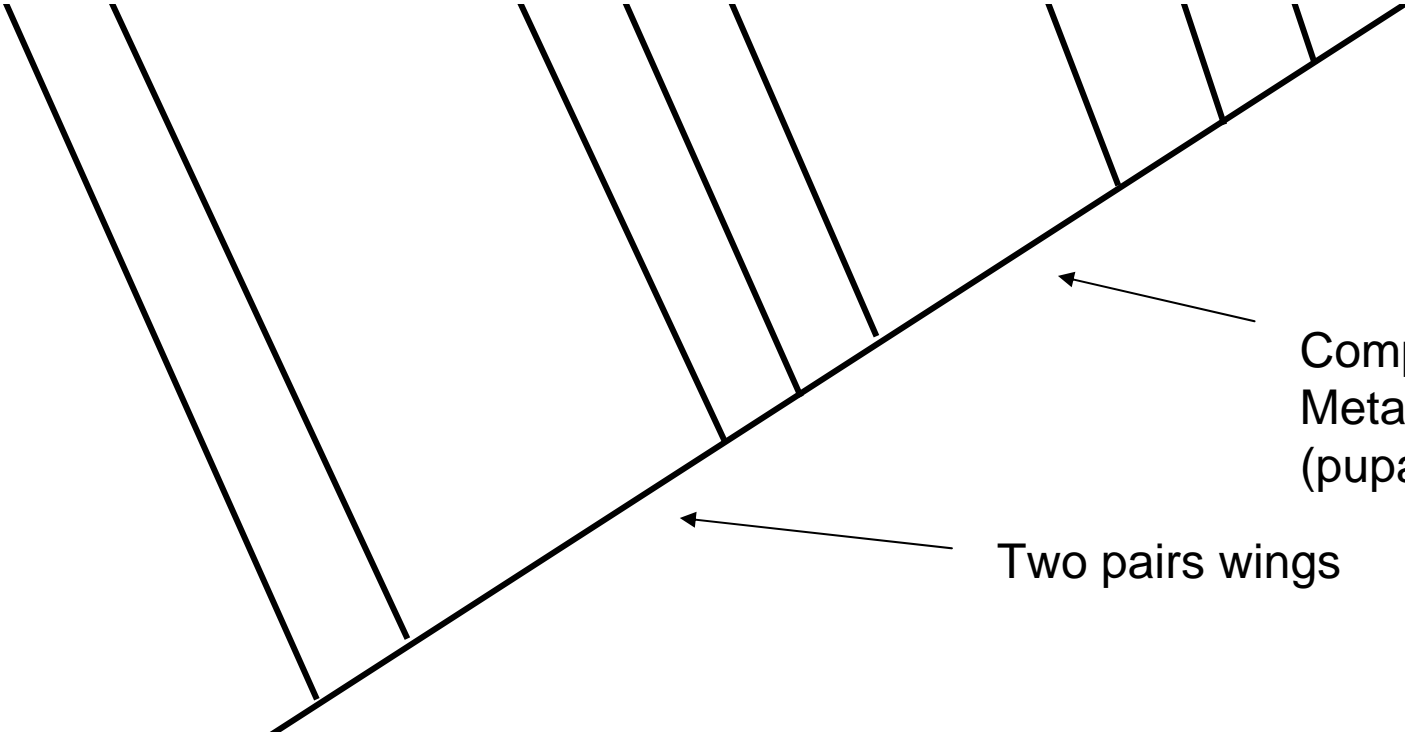
pea aphid

fly

Wingless insects

Hemimetabola

Holometabola



Complete Metamorphosis (pupa)

Two pairs wings

Holometabola

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e.g. beetle

honey bee

butterfly

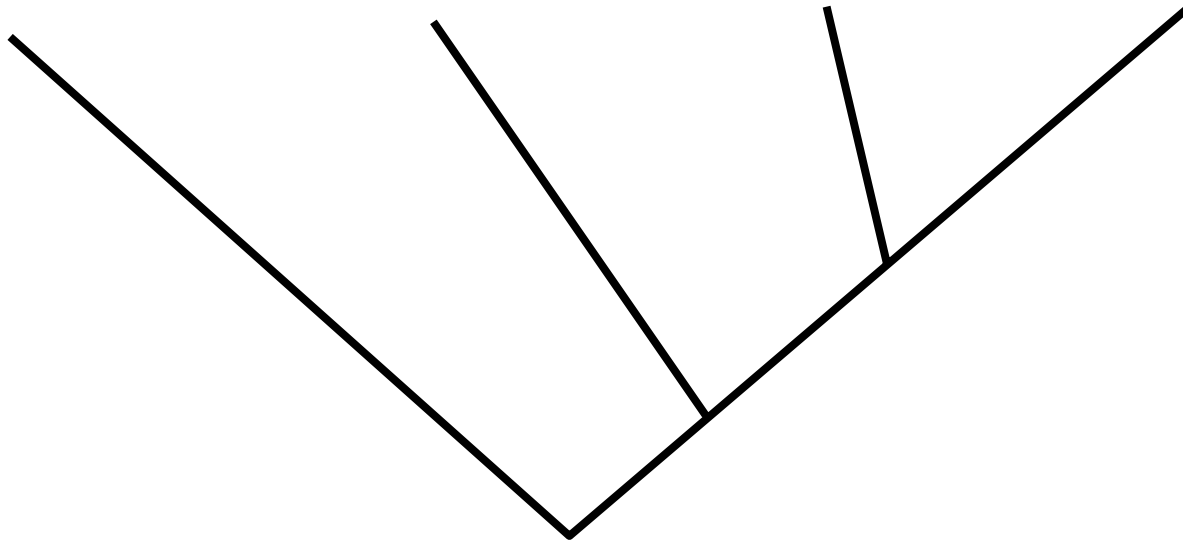
fly

Coleoptera

Hymenoptera

Lepidoptera

Diptera



Holometabola

Pictures from here removed for copyright reasons

e.g. beetle

honey bee

butterfly

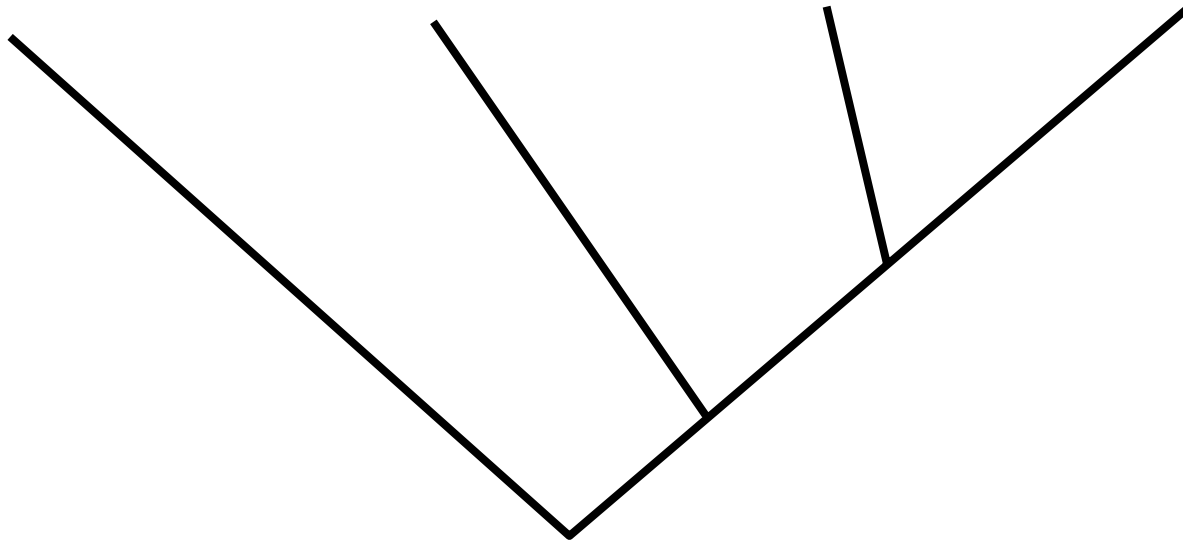
fly

Coleoptera

Hymenoptera

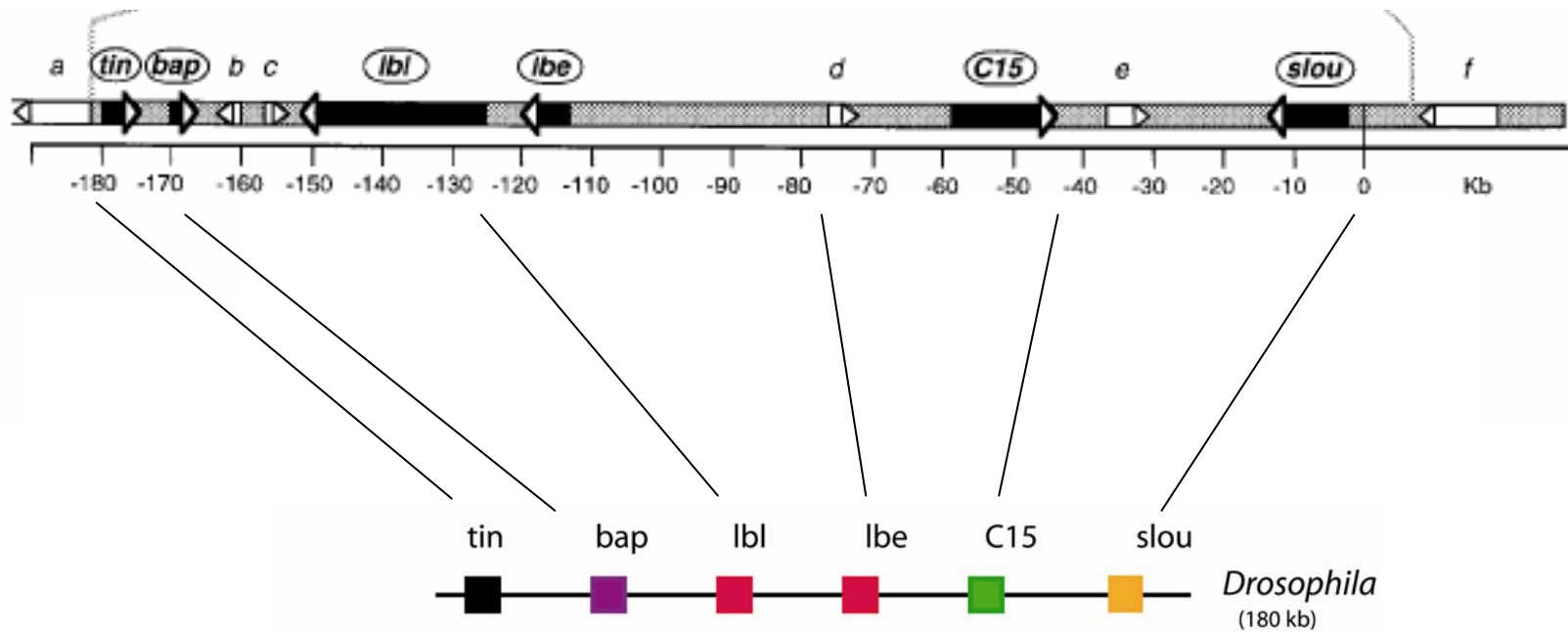
Lepidoptera

Diptera

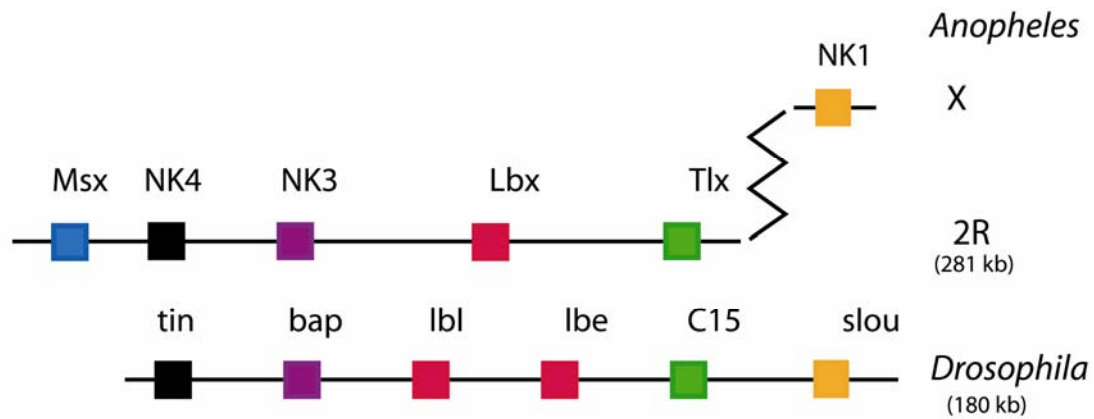


Now for an example of how comparative genomics
can be used to understand a set
of *Drosophila* genes

An unusual homeobox gene cluster in *Drosophila* : the NK-like genes



Gene name	Gene family	Expression/function
<i>tinman</i>	NK4 ■	all mesoderm; later dorsal muscle (e.g. heart)
<i>bagpipe</i>	NK3 ■	visceral mesoderm (e.g. around midgut)
<i>ladybird</i>	Lbx ■	heart cell fate; segmental border muscles
<i>C15</i>	Tlx ■	alary muscles
<i>slouch</i>	NK1 ■	somatic muscle: differentiation of muscle fibres



These fly genes have human orthologues;
therefore, it is an ancient gene cluster

Drosophila homeobox

Human homeobox

NK4 (tinman)

~ NKX-2.5, -2.6, -2.3



NK3 (bagpipe)

NKX-3.1, -3.2



lbe & lbl

LBX-1, -2



C15 (93Bal)

TLX-1, -2, -3

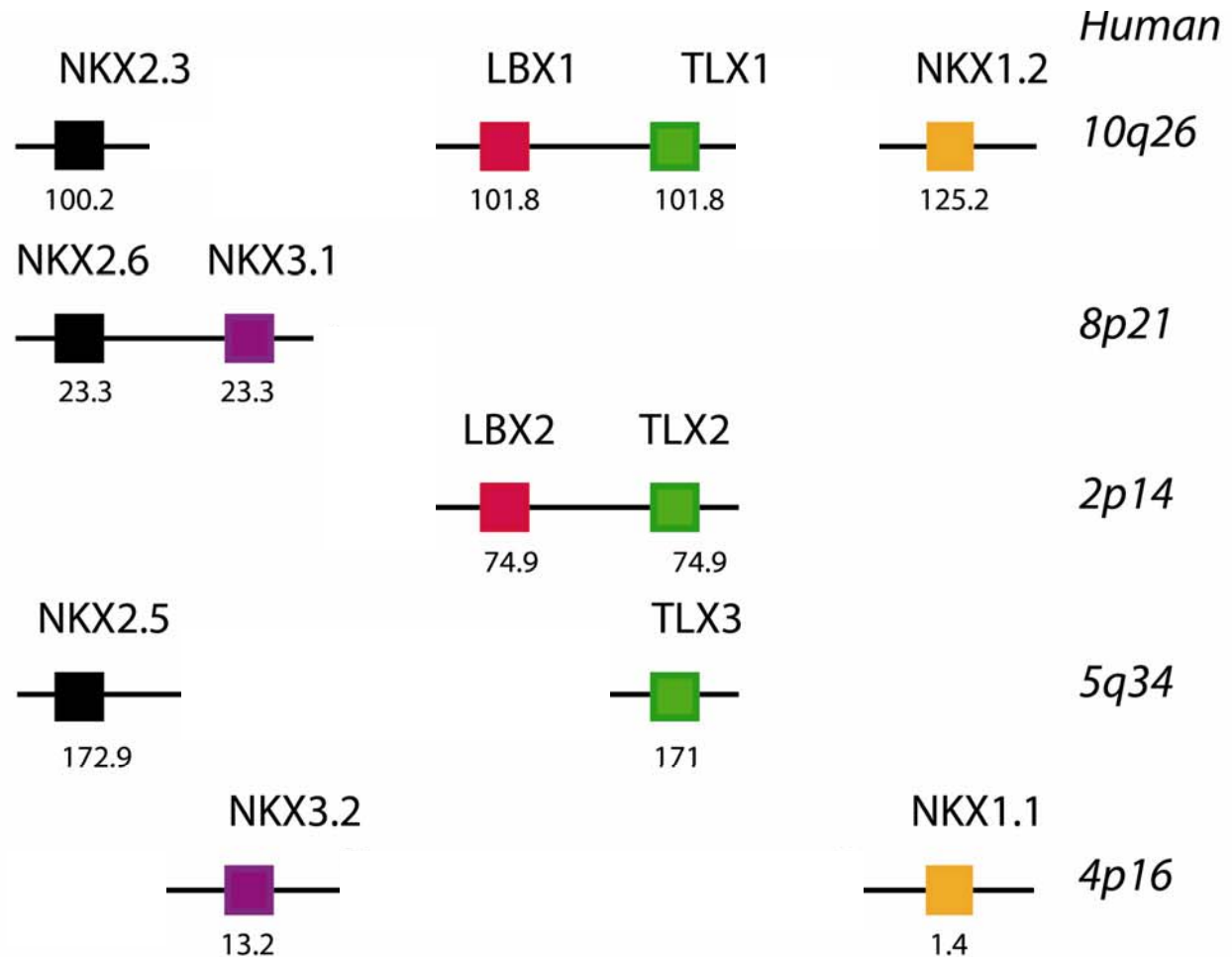


slouch (S59)

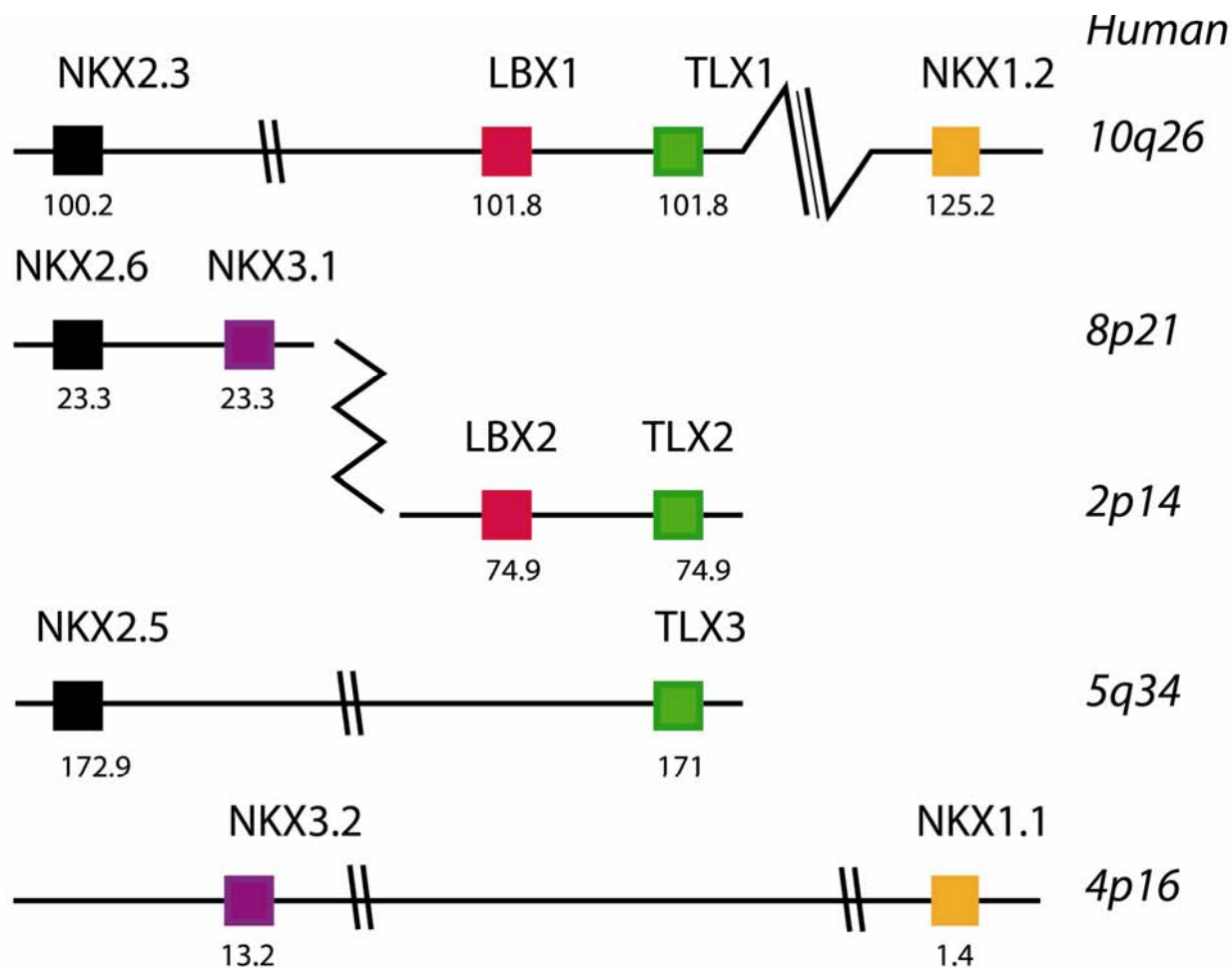
NKX-1.1, -1.2



'NK' class homeobox genes are scattered on 9 locations on 5 human chromosomes

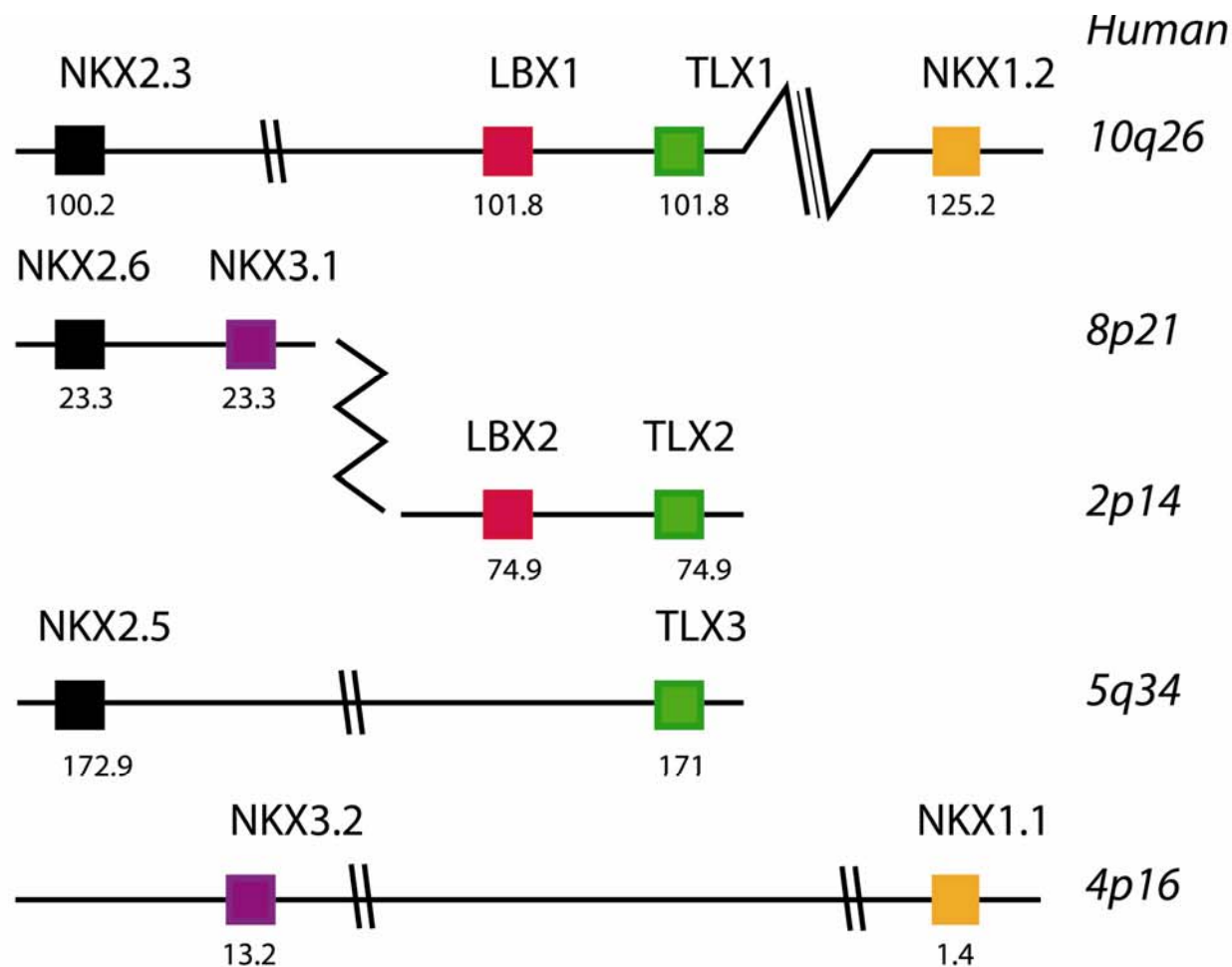


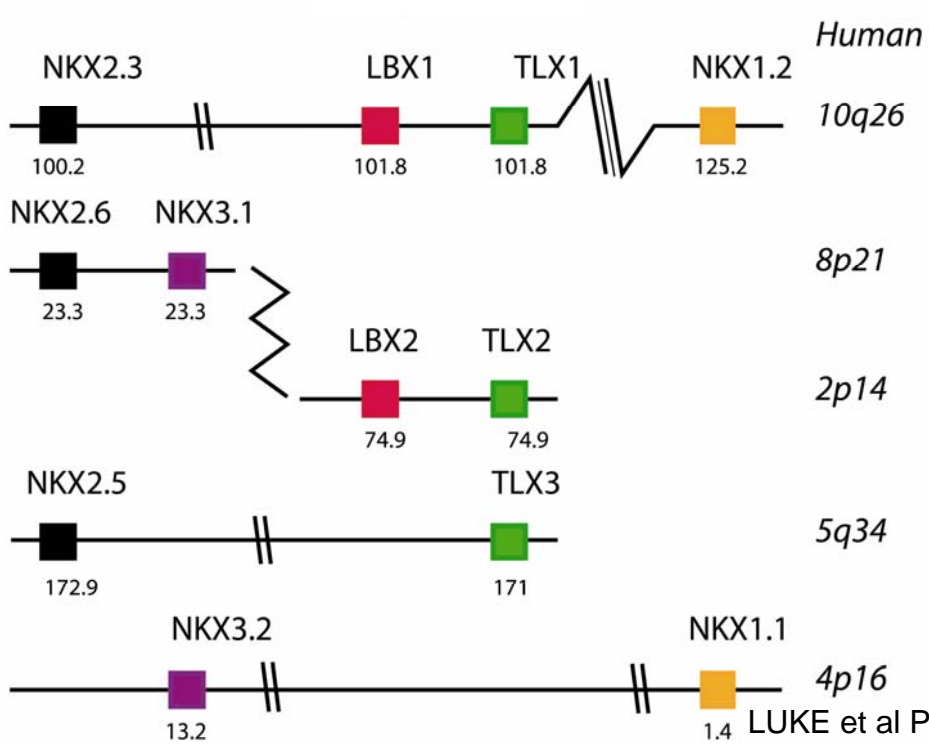
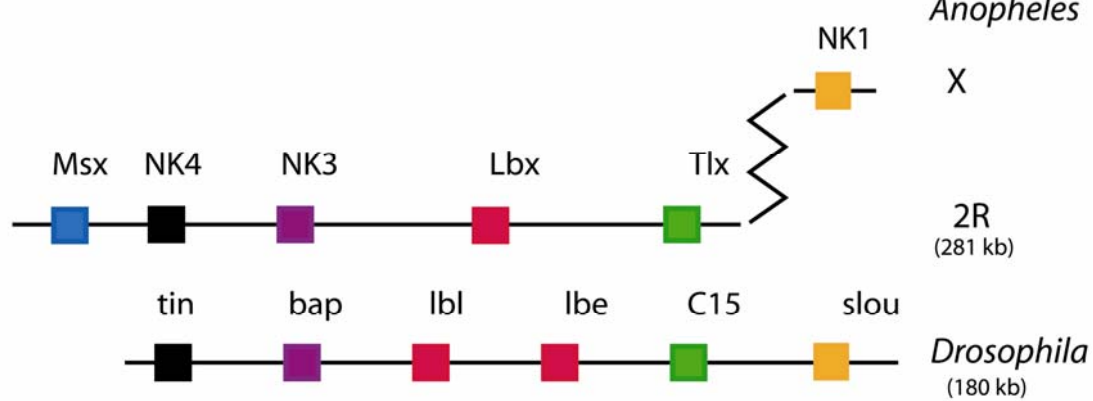
Maybe they are remnants of 4 gene clusters that have split and dispersed ?

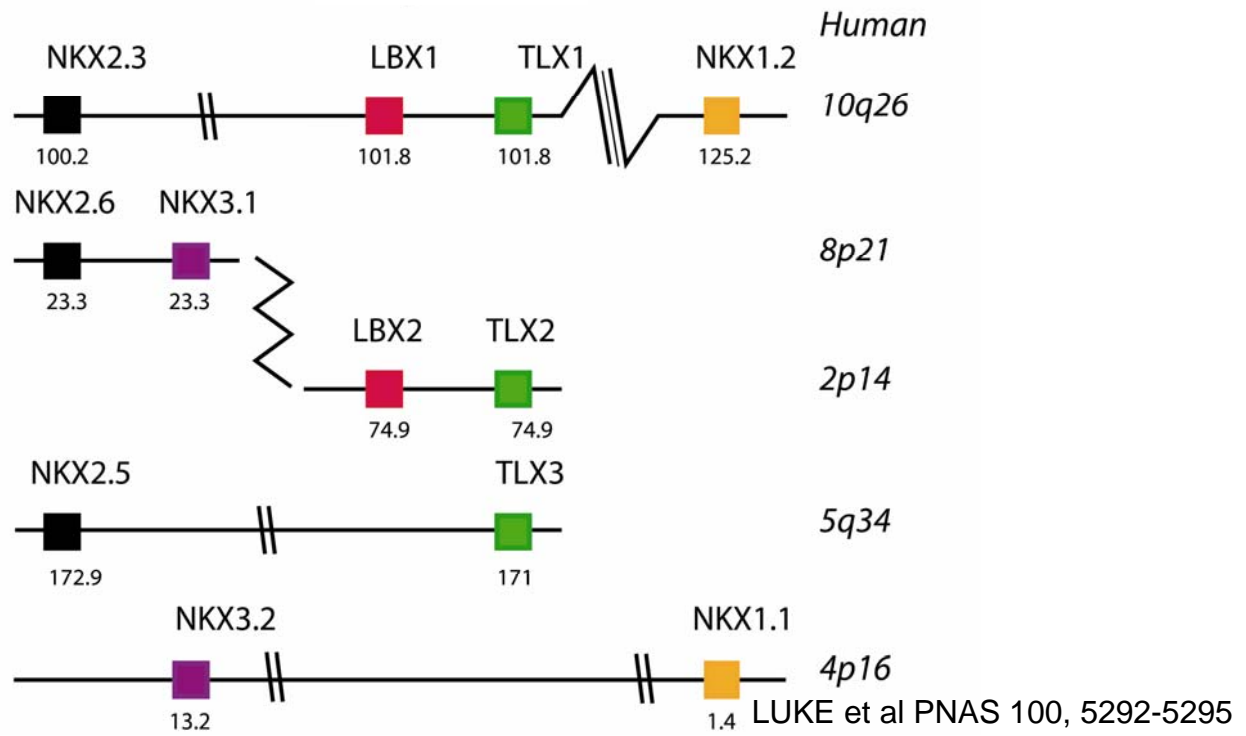
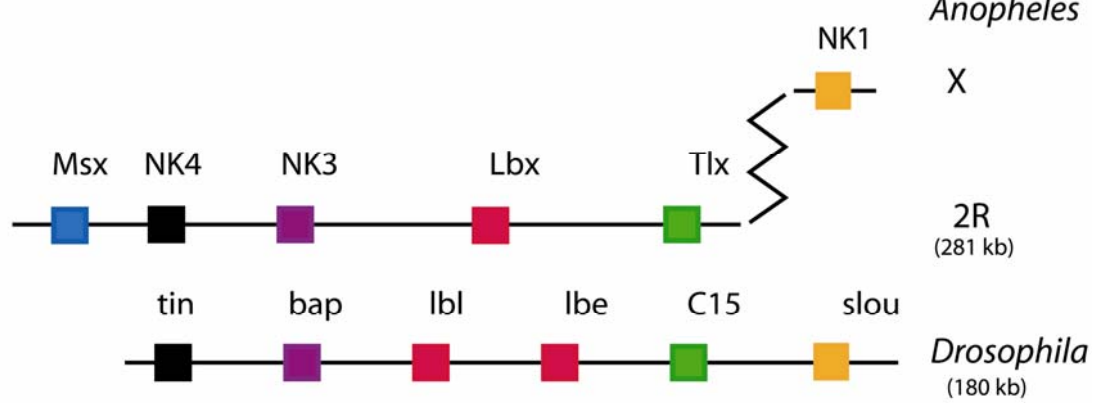




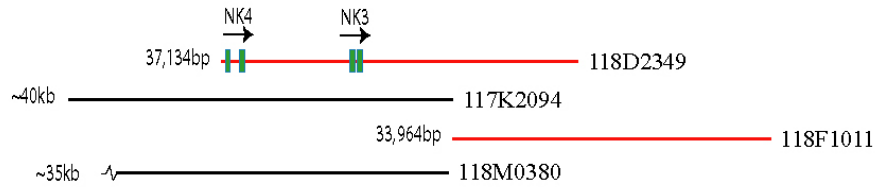
NKL



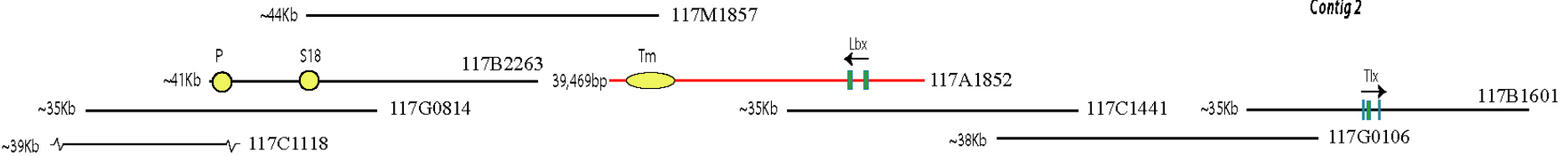




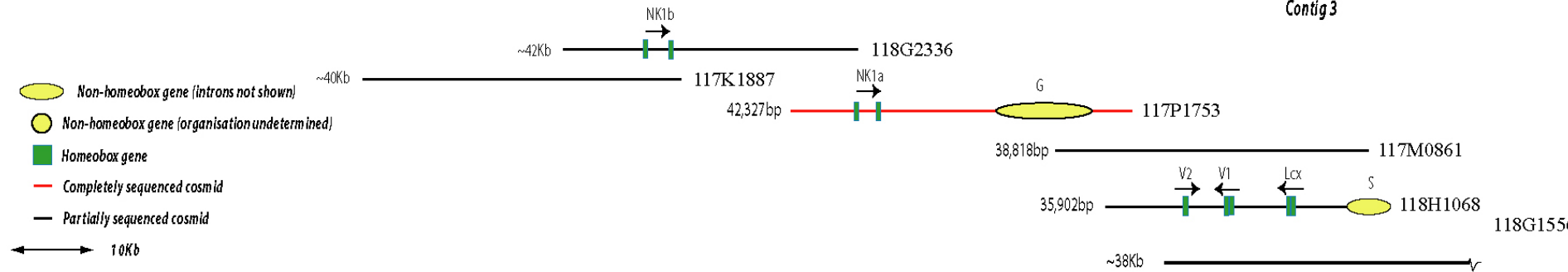
Contig 1



Contig 2



Contig 3



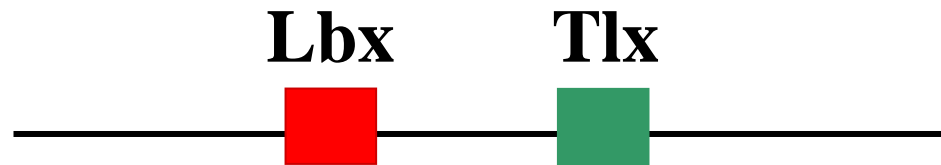
- Non-homeobox gene (introns not shown)
 - Non-homeobox gene (organisation undetermined)
 - Homeobox gene
 - Completely sequenced cosmid
 - Partially sequenced cosmid
- ←→ 10Kb



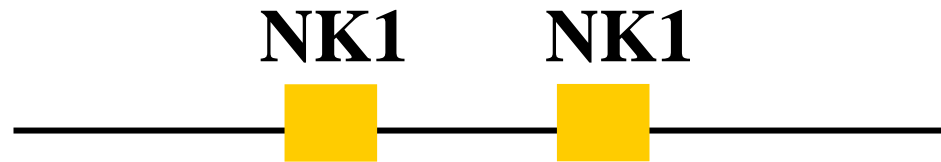
- Scaffold 1



- Scaffold 2

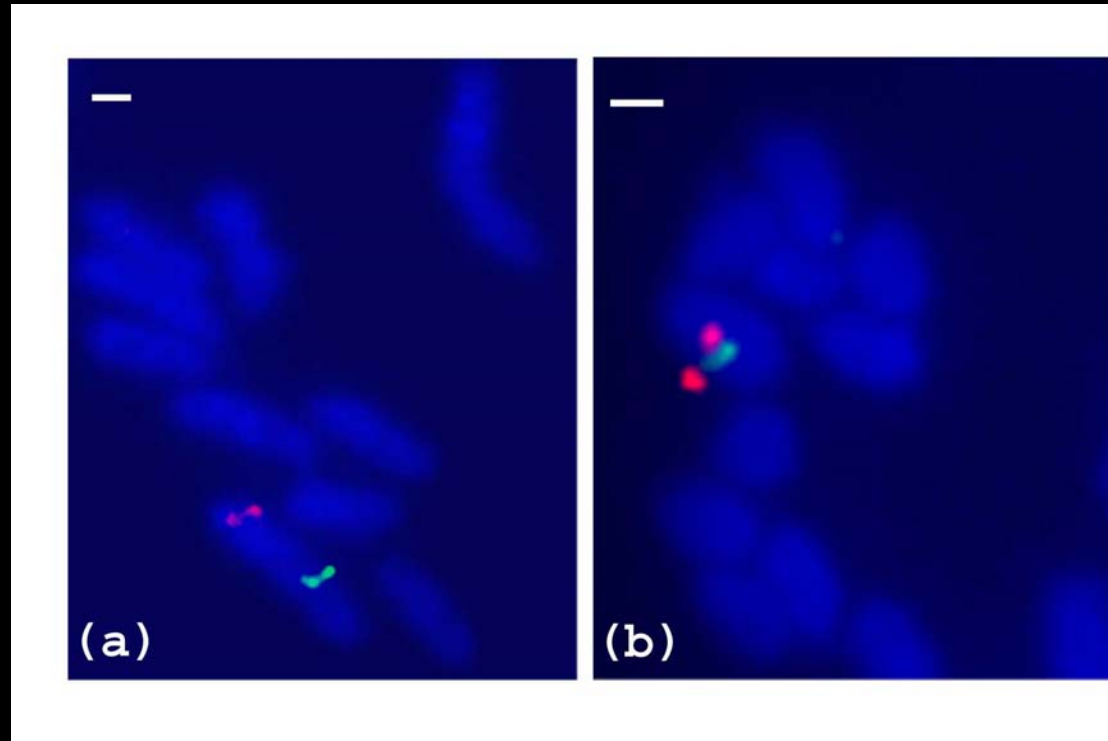


- Scaffold 3



DOUBLE PROBES

- a. Contig 1, **contig 2**
- b. Contig 3, **contig 2**

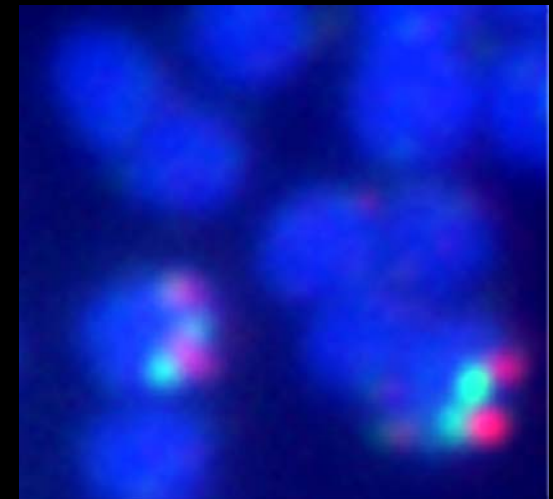


TRIPLE PROBE

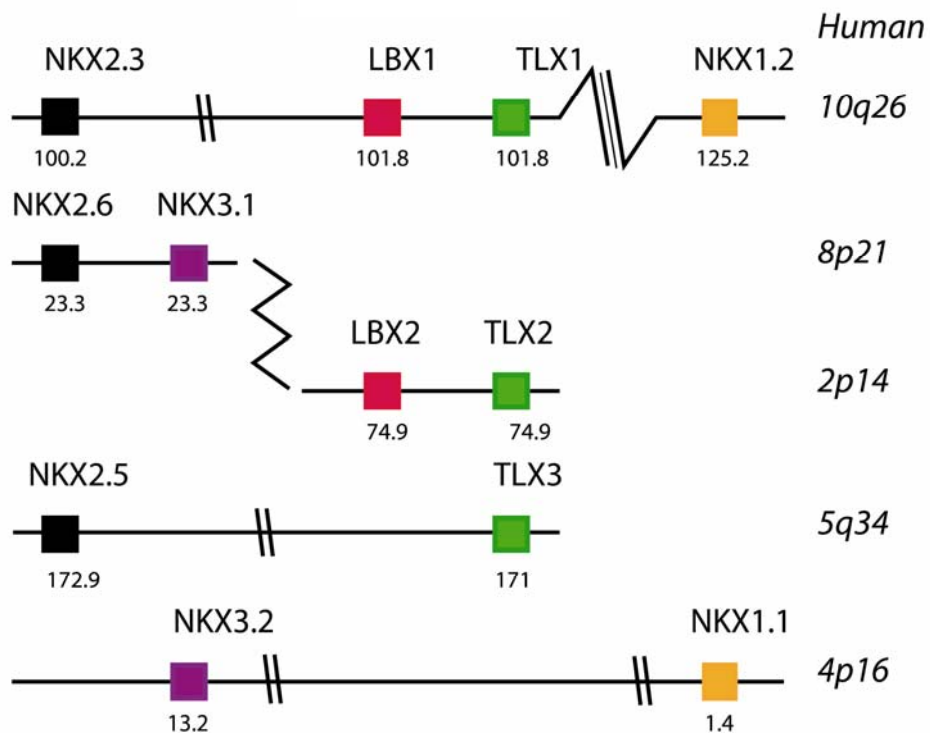
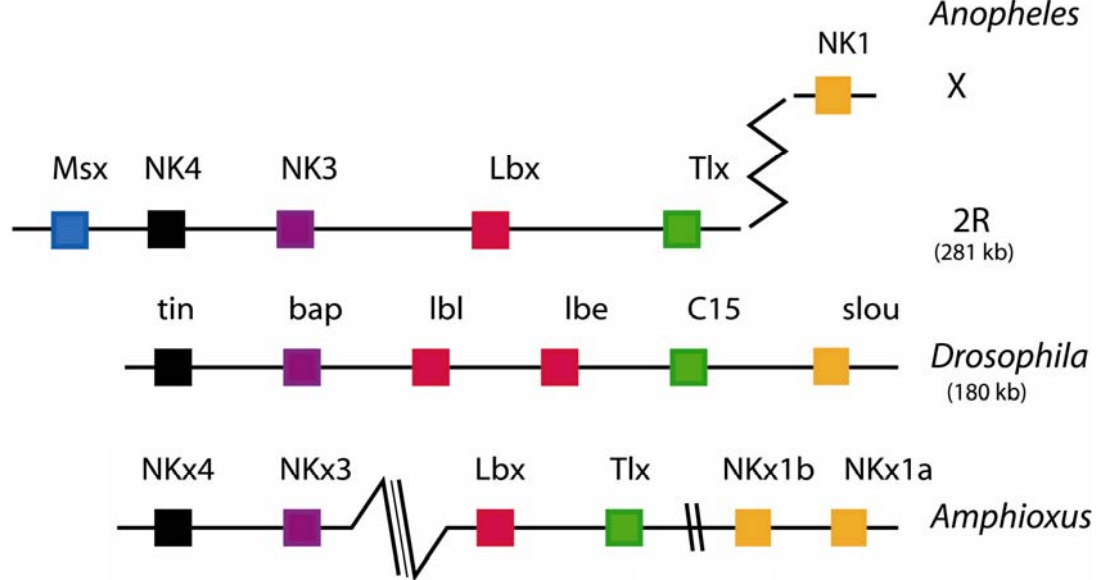
Contig 1 (green) **and** contig 2 **vs** contig 3 (red)



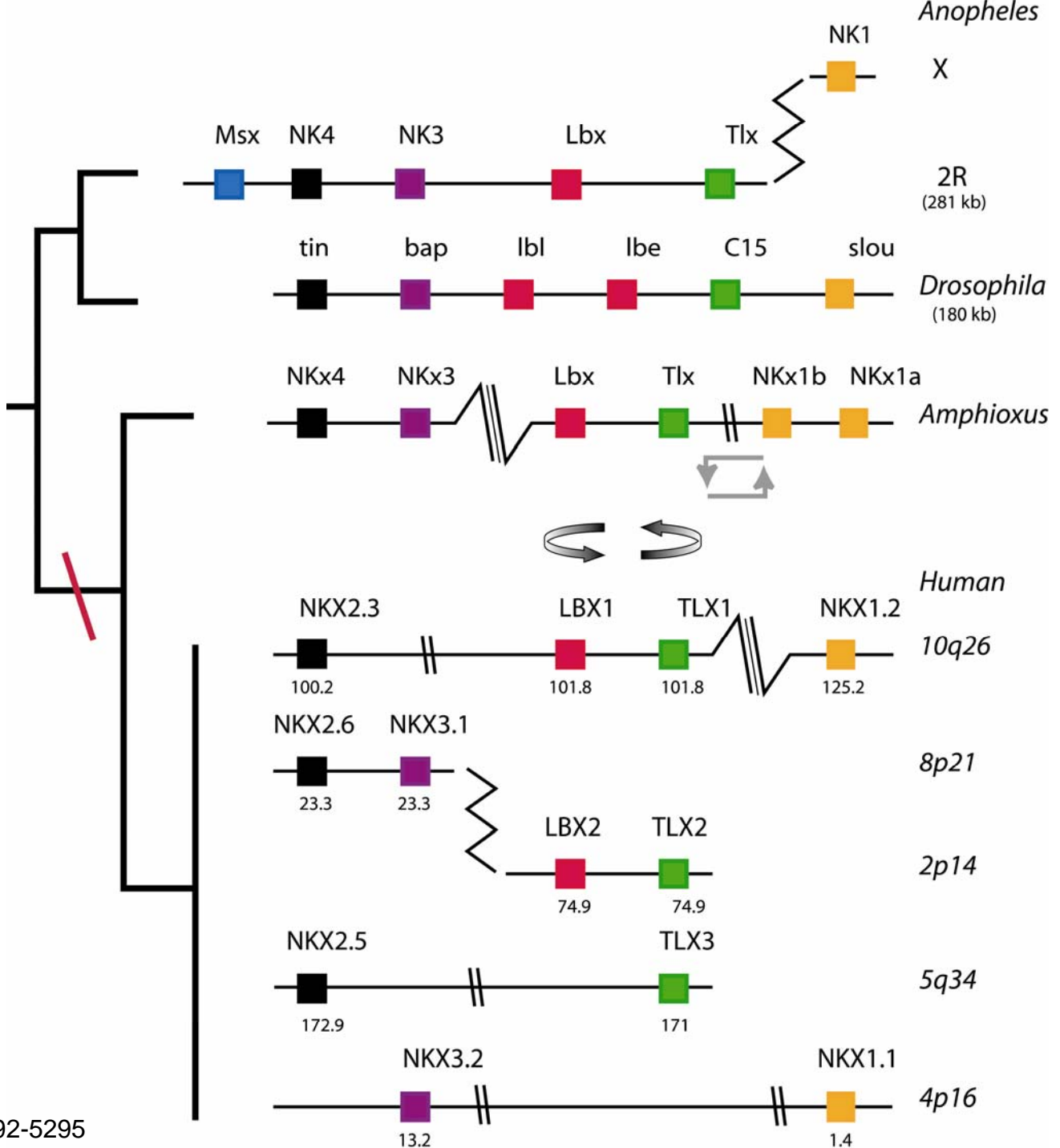
Filipe CASTRO



LUKE et al PNAS 100, 5292-5295



Ancient NK homeobox gene cluster



Conclusions

Dobzhansky “*Nothing in biology makes sense except in the light of evolution*”

Holland “*Nothing in genomics makes sense except in the light of evolution*”

- *Drosophila* is a metazoan (animal) – diversity of metazoan genomes
- *Drosophila* is a bilaterian – diversity of bilaterian genomes
- *Drosophila* is an ecdysozoan – insect and nematode genomes
- *Drosophila* is an arthropod – not very well sampled
- *Drosophila* is an insect – Holometabola well sampled, rest not

- Phylogeny plus genome sequences allows insight into genome evolution