

# Chapter 1

## Introduction, History of molecular Evolution

Welcome to the 2011 class

Chau-Ti Ting

ctting@ntu.edu.tw



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## Course description:

This course is to introduce the fundamentals and methodology on research of molecular evolution. The topics include evolutionary changes and patterns of sequences and methods for phylogenetic analyses. Topics on recent advances of molecular evolution studies will be offered at the end, on phylogenomics, horizontal gene transfer, and genome evolution. Students are highly recommended to take evolution, genetics or equivalent subjects before taking this course. There are three laboratory practices for students being familiar with data retrieving and analyses. Several homework assignments will be distributed during the semester, which require substantial extra work time after the class.



## Lecture

Wednesday 10:20~12:10, Life Science 419

## Instructors

Chau-Ti Ting ( 丁照棣 )

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Phone: 3366 2522

Jer-Ming Hu ( 胡哲明 )

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Phone: 3366 2472



**Course website:**

<http://ceiba.ntu.edu.tw/1002molevo>

**Official language:** English

**Prerequisite:** Biology; Evolutionary Biology; Genetics or equivalent preferred

**Office hour:** Wed 12:00-13:00, or with appointments



# Laboratory practice

We will have computers available to use on the class, but you are welcome to bring your own laptop computer. In the later case, you should download the corresponding softwares before the class. You are welcome to use your own data for the homework assignments.



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## Grading

Midterm exam 25%

Final exam 30%

Homework 35%

Classroom performance 10%



# Textbook

*There is no required textbook for this class. Most of the lectures are based on selected chapters from the following books.*



# Suggested reading

Graur, D. and W.-H. Li. 2000. *Fundamentals of Molecular Evolution*. 2<sup>nd</sup> ed., Sinauer Assoc., Sunderland, MA, USA.

Hall, B. G. 2011. Phylogenetic trees made easy: a how-to manual, 4<sup>th</sup> ed. Sinauer Assoc., Inc., Sunderland, MA, USA.

Li, W.-H. 1997. *Molecular evolution*. Sinauer Associates, Sunderland, MA, USA.

Lynch, M. 2007. *The origins of genome architecture*. Sinauer Associates, Sunderland, MA, USA.

Fundamentals of Molecular Evolution  
by Dan Graur and Wen-Hsiung Li, 2nd Edition (1999,  
Sinauer)

<http://www.sinauer.com/detail.php?id=266>



Hall, B. G. 2011. Phylogenetic trees made easy: a how-to manual, 4<sup>th</sup> ed. Sinauer Assoc., Inc., Sunderland, MA, USA.

<http://www.sinauer.com/detail.php?id=6069>



Li, W.-H. 1997. *Molecular evolution*. Sinauer Associates, Sunderland, MA, USA.

[http://books.google.com.tw/books/about/Molecular\\_evolution.html?id=2mPGQgAACAAJ&redir\\_esc=y](http://books.google.com.tw/books/about/Molecular_evolution.html?id=2mPGQgAACAAJ&redir_esc=y)



Lynch, M. 2007. The origins of genome architecture. Sinauer Associates, Sunderland, MA, USA.

<http://www.sinauer.com/detail.php?id=4843>



Molecular Evolution and Phylogenetics  
by Masatoshi Nei, Sudhir Kumar  
Oxford University Press (2000)

<http://www.oup.com/us/catalog/general/subject/LifeSciences/EvolutionaryBiology/?view=usa&isbn=0195135855>

Molecular Evolution : A Phylogenetic Approach  
by Roderic D. M. Page, Edward C. Holmes  
Blackwell Science Inc. (1998)  
<http://as.wiley.com/WileyCDA/WileyTitle/productCd-0865428891.html>

Integrated Molecular Evolution by  
Scott O. Rogers (Jul 27, 2011)

[http://www.crcpress.com/product/isbn/9781439819951;jsessionid=43hickpiklcWh5SL-z17dQ\\*\\*](http://www.crcpress.com/product/isbn/9781439819951;jsessionid=43hickpiklcWh5SL-z17dQ**)



## More reference book

Bioinformatics and Molecular Evolution by Paul Higgs  
and Teresa Attwood (Wiley-Blackwell, 2005)

\*Computational Molecular Evolution by Ziheng Yang  
(Oxford University Press, 2006)

\*Inferring Phylogenies (Paperback) by Joseph Felsenstein  
(Sinauer, 2003)

\*Statistical Methods in Molecular Evolution by Rasmus  
Nielsen (Springer, 2005)

The Phylogenetic Handbook: A Practical Approach to DNA  
and Protein Phylogeny Ed. by Marco Salemi and  
Anne-Mieke Vandamme (Cambridge University  
Press, 2003)



## Journals

Cladistics (

[http://www.blackwell-synergy.com/toc/cla/21/3;jsessionid=bfrbf8\\_Qdkn8](http://www.blackwell-synergy.com/toc/cla/21/3;jsessionid=bfrbf8_Qdkn8))

BMC Evol Biol (<http://www.biomedcentral.com/bmcevolbiol/>)

Evolution (<http://www.blackwell-synergy.com/loi/EVO>)

Evol Dev (<http://www.blackwell-synergy.com/loi/ede>)

Evol Bioinformatics (

[http://la-press.com/journal.php?journal\\_id=17&issue\\_id=103](http://la-press.com/journal.php?journal_id=17&issue_id=103))

Genome Biology (<http://genomebiology.com/home/>)

J Evol Biol ([http://www.blackwellpublishing.com/jeb\\_enhanced/](http://www.blackwellpublishing.com/jeb_enhanced/))

J Mol Evol (<http://www.springer.com/life+sci/cell+biology/journal/239>)

Mol Biol Evol (<http://mbe.oxfordjournals.org/>)

Mol Ecol (<http://www.blackwell-synergy.com/loi/MEC>)

Mol Phyl Evol (<http://www.sciencedirect.com/science/journal/10557903>)

## Related courses and workshops

### **NTU campus**

Mathematical Molecular Evolution (221 U4730)

Molecular Evolution Laboratory (B44 U1630)

Population Genetics (B44 U1570)

Seminar in Molecular Evolution (B41 M0310)

Summer course on molecular phylogeny

Workshop on Computational Molecular Evolution (

<http://homepage.ntu.edu.tw/~ctting/CMB.html>)

### **Elsewhere**

Summer Institute in Statistical Genetics

<http://www.biostat.washington.edu/node/1020>)

Workshop on Molecular Evolution (

<http://workshop.molecularevolution.org/mbl/>



# *Molecular Evolution*

The evolution of macromolecules

- The rates and patterns of changes in the genetic material and its encoded products during evolutionary time
- The mechanisms responsible for such changes

Molecular phylogeny

- The reconstruction of the evolutionary history of genes and organisms

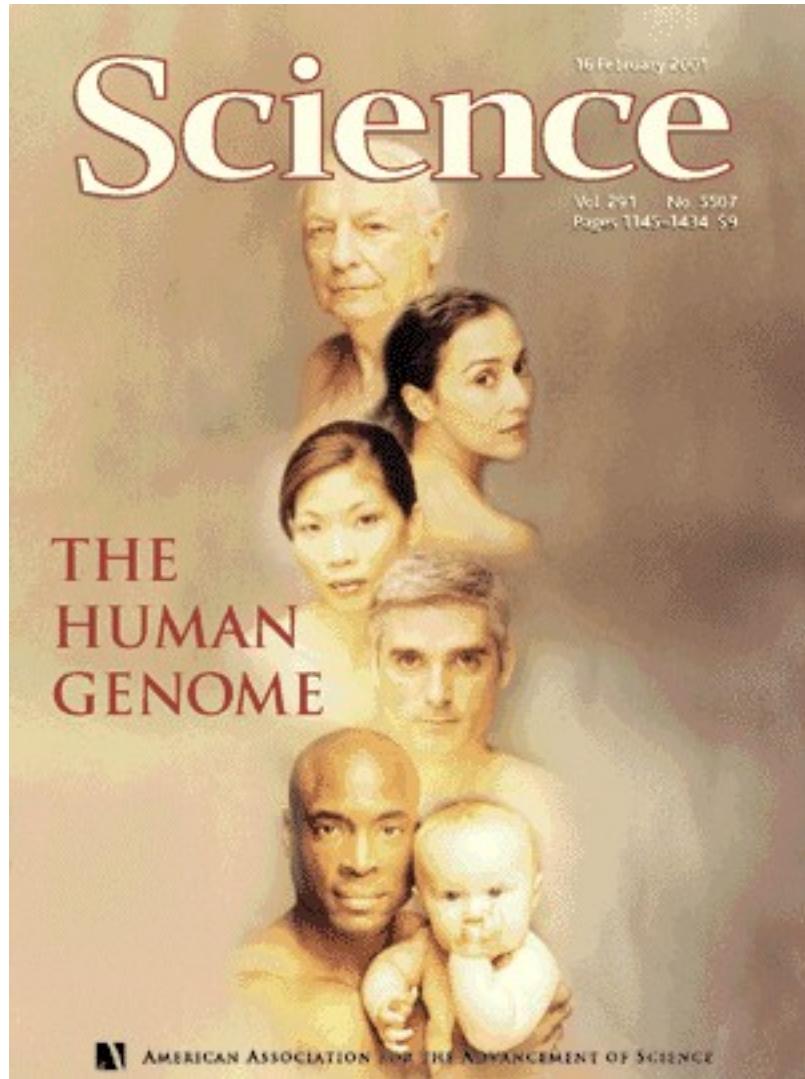
Source: M. Prakash

2007. Encyclopaedia of Gene Evolution Vol. 2, Molecular Genetics, p. 42. Discovery Publishing House. India.

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OpenCourseWare

臺大開放式課程



 Science /AAAS

Source: 2001. *Science*. Vol. 291. Issue 5507.

<http://www.sciencemag.org/content/291/5507.cover-expansion>

## GENETIC ORIGINS

*The study of human evolution begins with your DNA.*



 Carolina Biological Supply Company

Source: Carolina Biological Supply Company

<http://www.geneticorigins.org/>

# 1904

Nuttall conducted precipitin tests of serum proteins to infer the phylogenetic relationship among various groups of animals.

 Source: M. Prakash  
2007. Encyclopaedia of Gene Evolution Vol. 2, Molecular Genetics, p. 42. Discovery Publishing House. India.

George H. F. Nuttall  
1904. Blood immunity and blood relationship. University press. Cambridge, England.

<http://www.biodiversitylibrary.org/title/8418#page/7/mode/1up>

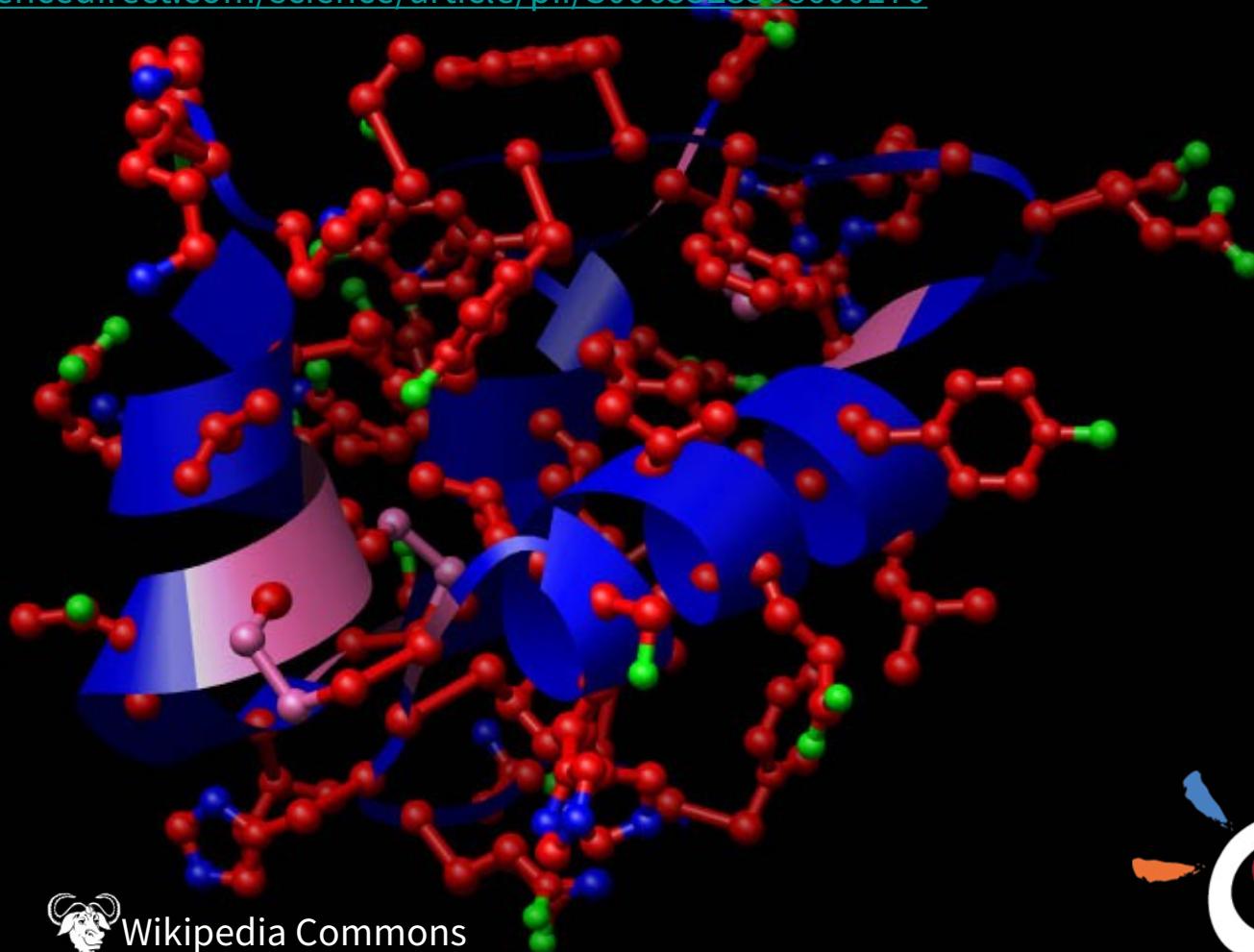


# 1952

## Sanger and colleagues complete first protein sequence – insulin

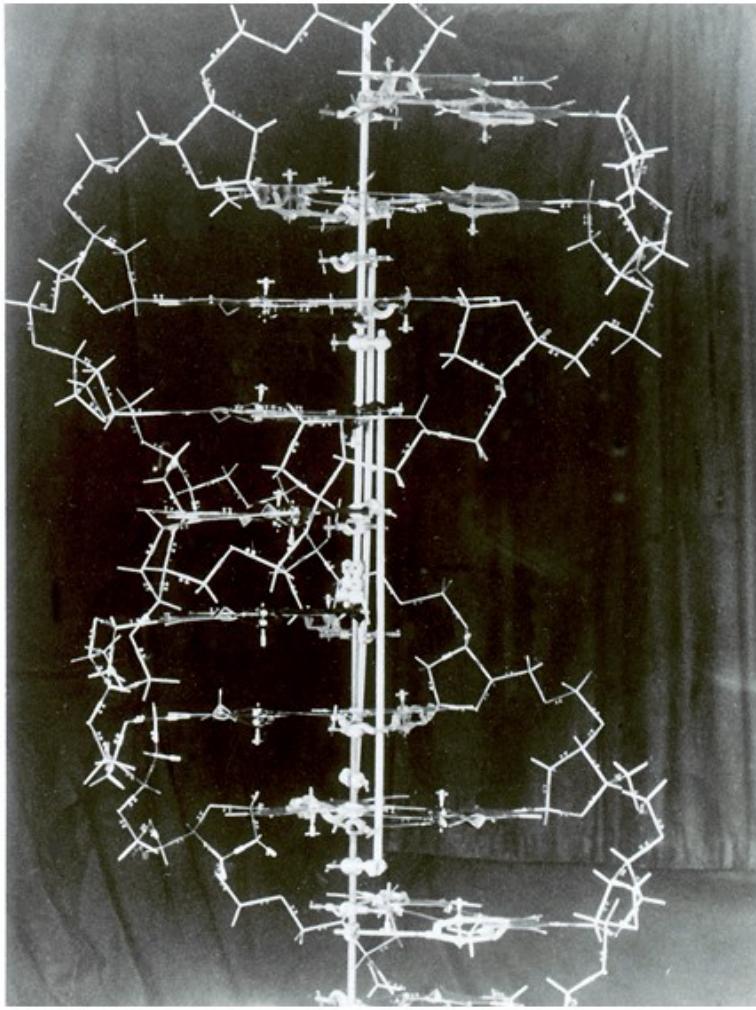
Sanger, F. 1952. The arrangement of amino acids in proteins. Advances in Protein Chemistry. 7: p.1-66.

<http://www.sciencedirect.com/science/article/pii/S0065323308600170>



Wikipedia Commons

# 1953



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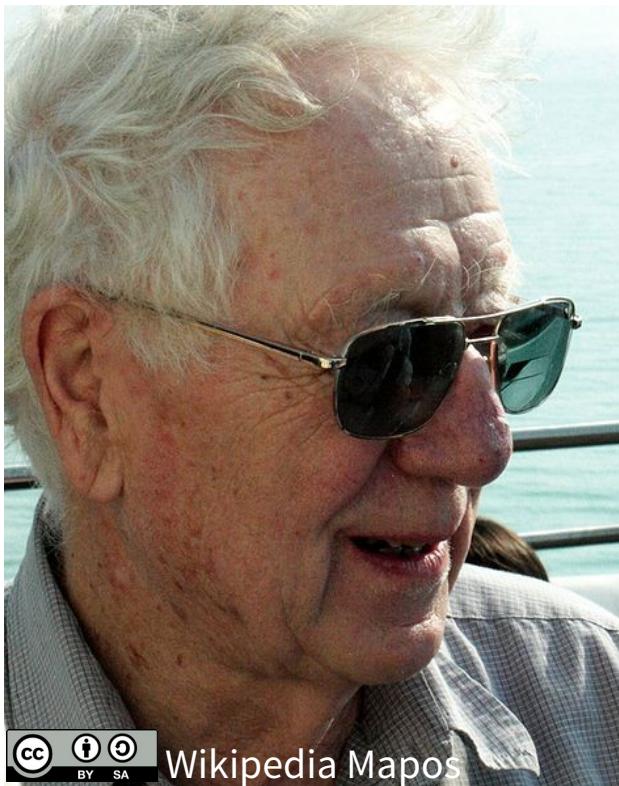
Watson and Crick in 1953. Cold Spring Harbor Laboratory

# 1955

Smithies uses starch-gel electrophoresis to identify protein polymorphisms

 John C. Avise

<http://www.ncbi.nlm.nih.gov/pmc/articles/PMC1215845/>



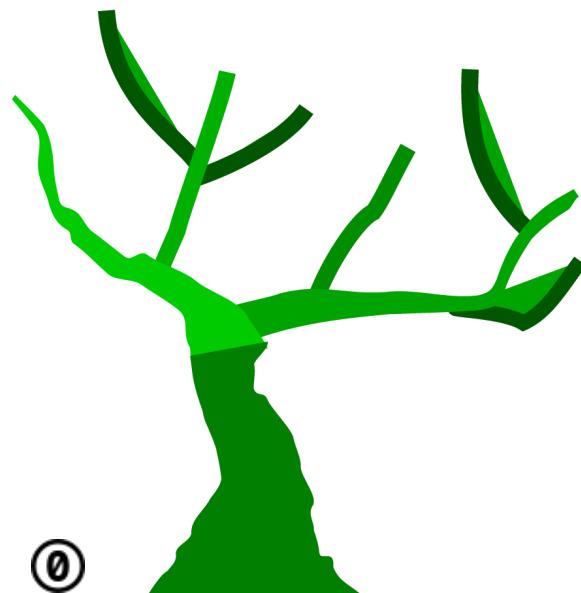
Wikipedia Mapos



# 1963

Margoliash determines amino acid sequences for cytochrome c in several taxa and generates the first phylogenetic tree for a specific gene product.

 John C. Avise <http://www.ncbi.nlm.nih.gov/pmc/articles/PMC221244/>

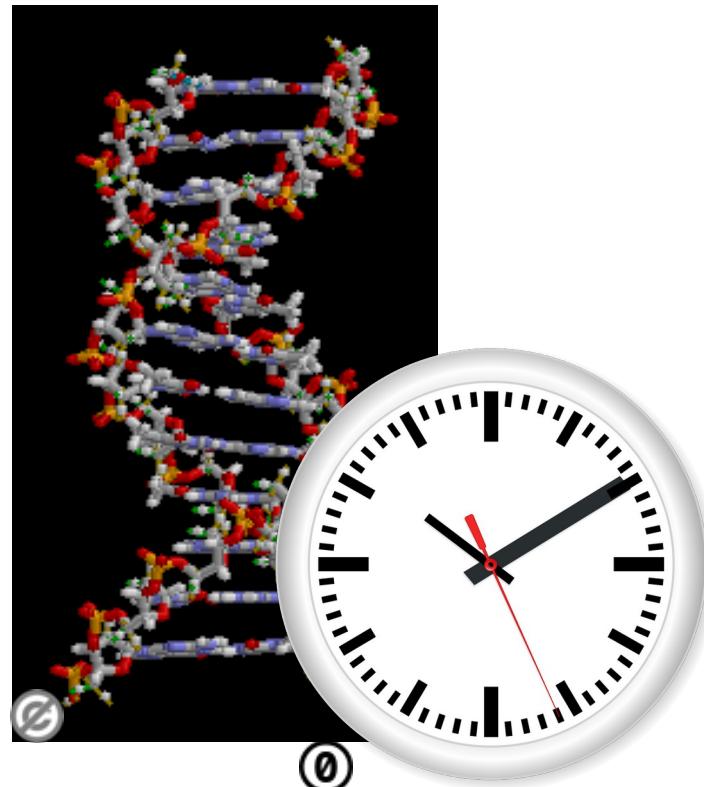


# 1965

Zuckerkandl and Pauling

- “molecular clock” hypothesis.

[http://en.wikipedia.org/wiki/History\\_of\\_molecular\\_evolution](http://en.wikipedia.org/wiki/History_of_molecular_evolution)



# 1966

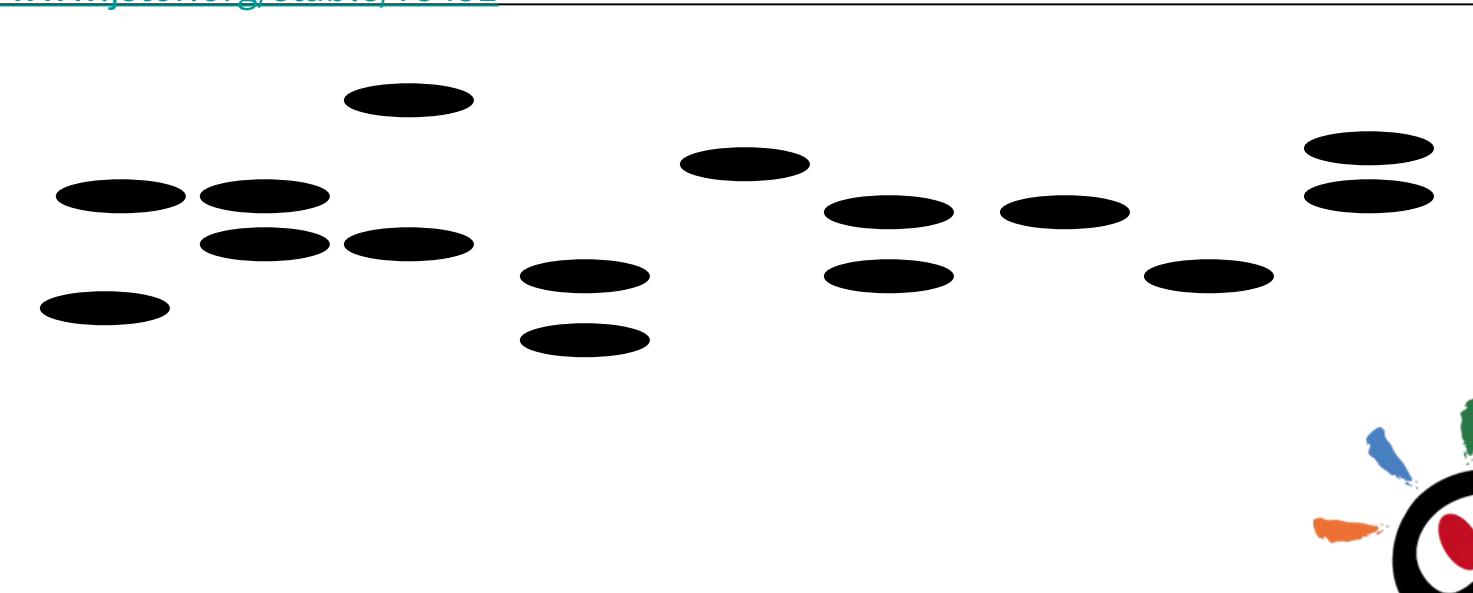
Harris, Lewontin and Hubby

- electrophoresis and histochemical enzyme stains
- examined level of genetic variability in natural populations.

[http://en.wikipedia.org/wiki/Richard\\_Lewontin](http://en.wikipedia.org/wiki/Richard_Lewontin)

<http://www.ncbi.nlm.nih.gov/pmc/articles/PMC1211186/?tool=pmcentrez>

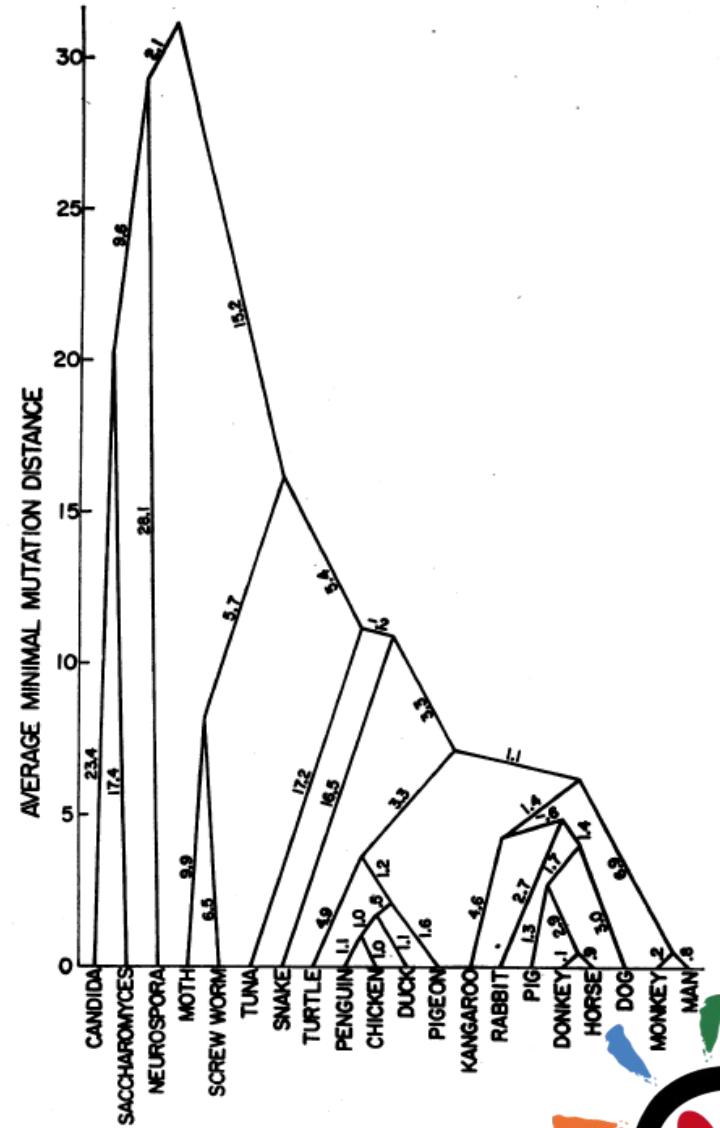
<http://www.jstor.org/stable/75451>



# 1967

Fitch and Margoliash  
•“Construction of phylogenetic trees”  
published in Science.

Walter M. Fitch and Emanuel Margoliash  
1967. Construction of phylogenetic trees.  
Science Vol. 155: p. 279-284



Source: Walter M. Fitch and Emanuel Margoliash  
1967. Construction of phylogenetic trees. Science Vol. 155: p.  
279-284  
<http://www.sciencemag.org/content/155/3760/279.full.pdf>

CC/NUMBER 27  
JULY 4, 1988

# This Week's Citation Classic®

Fitch W M & Margoliash E. Construction of phylogenetic trees.  
Science 155:279-84, 1967.

[University of Wisconsin Medical School, Madison, WI and Protein Section, Department of Molecular Biology, Abbott Laboratories, North Chicago, IL]

This paper provided a way to construct an evolutionary tree using a measure of genetic distance obtained from amino acid sequence differences interpreted in terms of the genetic code. [The SCI® indicates that this paper has been cited in over 670 publications.]

revealed that there was some slowness in the rate of sequence publication from his laboratory and that he had another 10 completed sequences. He offered to provide them to me. What a windfall! At the stroke of a conversational noon-hour clock, the sample size was doubled to 20, permitting us to present a tree spanning the largest part of the eukaryotic kingdom, and I got a coauthor whose prose was pellucid.

The genetic code was solved (except we then thought that the methionine and tryptophan codons were twofold degenerate), the minimum base differences were plugged into the

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Walter M. Fitch

Ahmanson Center for Biological Research  
University of Southern California  
Los Angeles, CA 90089-1481



Source: Walter M. Fitch. 1988. Citation Classic. No. 27.

<http://garfield.library.upenn.edu/classics.html>

<http://garfield.library.upenn.edu/classics1988/A1988N88820001.pdf>

# 1968

Kimura proposes that the majority of molecular changes in evolution are due to random drift of neutral or nearly neutral mutations.

[http://en.wikipedia.org/wiki/Motoo\\_Kimura#cite\\_note-Kimura68-1](http://en.wikipedia.org/wiki/Motoo_Kimura#cite_note-Kimura68-1)  
<http://www.nature.com/nature/journal/v217/n5129/abs/217624a0.html>



Source: John C. Avise

1994. Molecular Markers, Natural History and Evolution, p. 42. Chapman & Hall. New York, USA.

# 1969

King and Jukes

- “Non-Darwinian evolution”
- Also proposed neutral theory .

[http://en.wikipedia.org/wiki/Non-Darwinian\\_Evolution](http://en.wikipedia.org/wiki/Non-Darwinian_Evolution)

<http://www.sciencemag.org/content/164/3881/788>

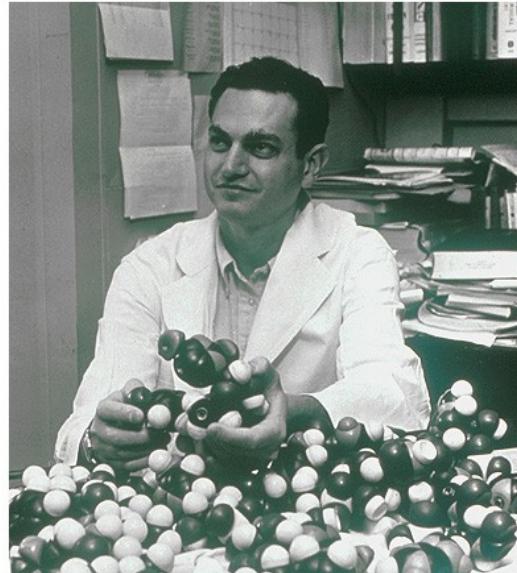


# 1961

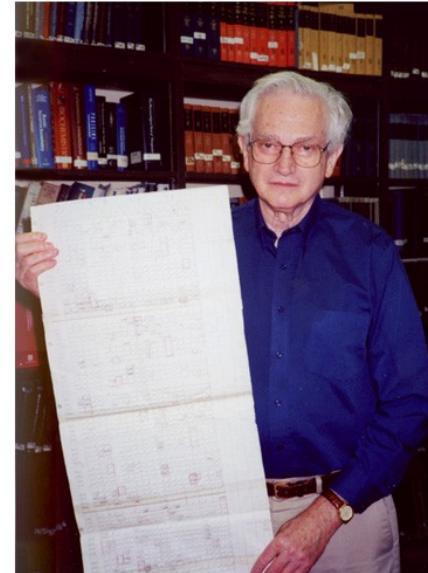
Marshall Nirenberg, a young biochemist at the National Institute of Arthritic and Metabolic Diseases, discovered the first "triplet"—a sequence of three bases of DNA that codes for one of the twenty amino acids that serve as the building blocks of proteins. Subsequently, within five years, the entire **genetic code** was deciphered.

Source: Genome News Network

[http://www.genomenewsnetwork.org/resources/timeline/1961\\_Nirenberg.php](http://www.genomenewsnetwork.org/resources/timeline/1961_Nirenberg.php)



Courtesy of the National Library of Medicine.  
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Courtesy of Dr. S. Chan, DNA Learning Center.  
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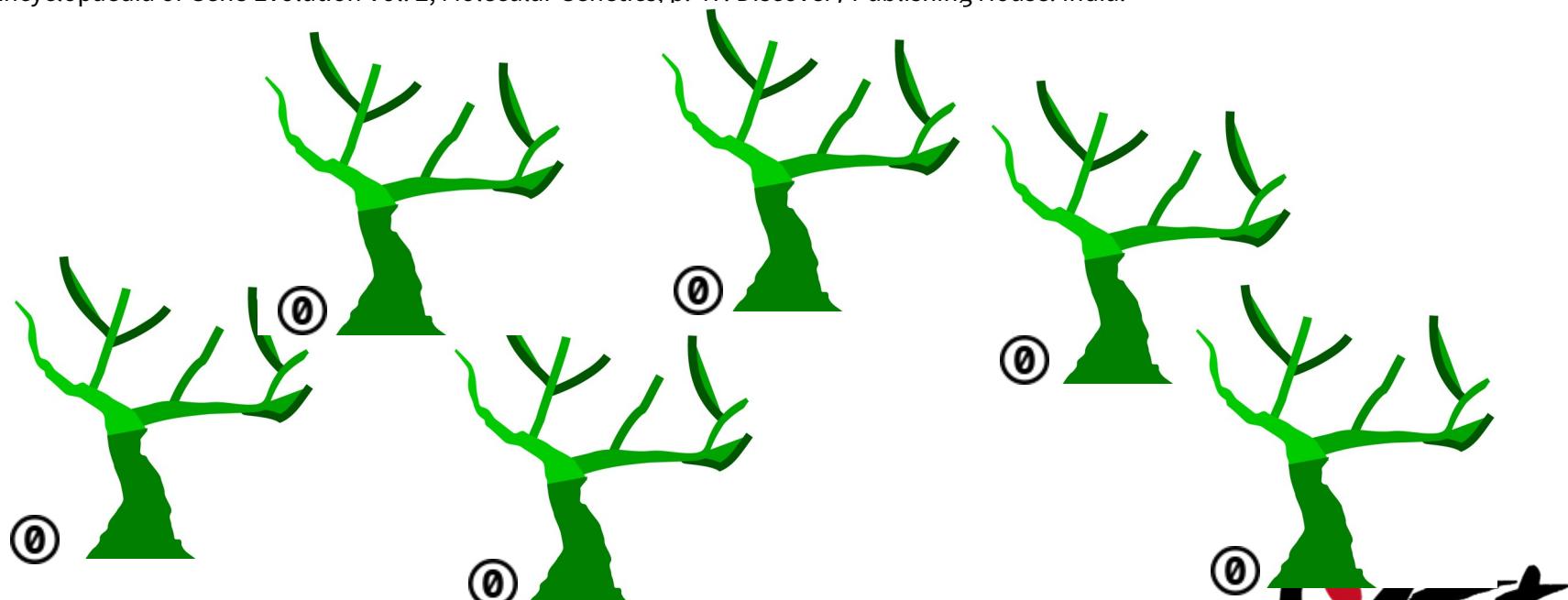
# 1972

Nei's genetic distance has greatly facilitated the study of evolutionary relationships among populations or closely related species.

[http://en.wikipedia.org/wiki/Masatoshi\\_Nei](http://en.wikipedia.org/wiki/Masatoshi_Nei)  
<http://www.jstor.org/stable/2459777>

Source: M. Prakash

2007. Encyclopaedia of Gene Evolution Vol. 2, Molecular Genetics, p. 47. Discovery Publishing House. India.



# 1977

Molecular biologists by the 1970s had deciphered the genetic code and could spell out the sequence of amino acids in proteins. But inability to easily read off the precise nucleotide sequences of DNA forestalled further advances in molecular genetics and all prospects of genetic engineering. Walter Gilbert (with graduate student Allan M. Maxam) and Frederick Sanger, in 1977, working separately in the United States and England, developed new techniques for rapid DNA sequencing.

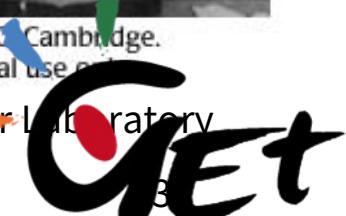
 Source: Genome News Network



Courtesy of Dr. F. Sanger, MRC Cambridge.  
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General Education TW  
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# 1981

Palmer and colleagues begin an important series  
papers utilizing cpDNA for phylogenetic  
 **constructions in plants**

Source: J. Palmer et al.

1994. Molecular Markers, Natural History and Evolution, p. 42. Chapman & Hall. New York, USA.

# 1984

Sibley and Ahlquist  
•DNA-DNA hybridization  
•primate evolution study.

[http://en.wikipedia.org/wiki/DNA-DNA\\_hybridization](http://en.wikipedia.org/wiki/DNA-DNA_hybridization)

<http://www.springerlink.com/content/g3020651ml56620/>





Science /AAAS

Source: 2001. Science. Vol. 291. Issue 5507.

<http://www.sciencemag.org/content/291/5507.cover-expansion>

## GENETIC ORIGINS

*The study of human evolution begins with your DNA.*



Carolina Biological Supply Company

Source: Carolina Biological Supply Company  
<http://www.geneticorigins.org/>

# Homework Assignment I

- 1) Write a paragraph about yourself (including name, background, and contact information) and your research interests.
- 2) What do you expect to learn from taking this course?

Please send your homework as a pdf document  
online by Feb. 28th



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P17. "The rates and patterns of changes in the genetic material and its encoded products during evolutionary time"		<p>M. Prakash          2007. Encyclopaedia of Gene Evolution Vol. 2, Molecular Genetics, p. 42. Discovery Publishing House. India.  <a href="http://books.google.com.tw/books?id=OoG4_y2Q9AEC&amp;printsec=frontcover&amp;hl=zh-TW#v=onepage&amp;q&amp;f=false">http://books.google.com.tw/books?id=OoG4_y2Q9AEC&amp;printsec=frontcover&amp;hl=zh-TW#v=onepage&amp;q&amp;f=false</a>          2012/02/21 visited  <i>It is used subject to the fair use doctrine of:</i>          • Taiwan Copyright Act Articles 52 &amp; 65</p>	P17
P17. "The mechanisms responsible for such changes"		<p>M. Prakash          2007. Encyclopaedia of Gene Evolution Vol. 2, Molecular Genetics, p. 42. Discovery Publishing House. India.  <a href="http://books.google.com.tw/books?id=OoG4_y2Q9AEC&amp;printsec=frontcover&amp;hl=zh-TW#v=onepage&amp;q&amp;f=false">http://books.google.com.tw/books?id=OoG4_y2Q9AEC&amp;printsec=frontcover&amp;hl=zh-TW#v=onepage&amp;q&amp;f=false</a>          2012/02/21 visited  <i>It is used subject to the fair use doctrine of:</i>          • Taiwan Copyright Act Articles 52 &amp; 65</p>	P17
P17. "The reconstruction of the evolutionary history of genes and organisms "		<p>M. Prakash          2007. Encyclopaedia of Gene Evolution Vol. 2, Molecular Genetics, p. 42. Discovery Publishing House. India.  <a href="http://books.google.com.tw/books?id=OoG4_y2Q9AEC&amp;printsec=frontcover&amp;hl=zh-TW#v=onepage&amp;q&amp;f=false">http://books.google.com.tw/books?id=OoG4_y2Q9AEC&amp;printsec=frontcover&amp;hl=zh-TW#v=onepage&amp;q&amp;f=false</a>          2012/02/21 visited  <i>It is used subject to the fair use doctrine of:</i>          • Taiwan Copyright Act Articles 52 &amp; 65</p>	P17
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		<p>Watson and Crick in 1953.Cold Spring Harbor Laboratory  <a href="http://www.dnalc.org/view/16430-Gallery-19-DNA-model-1953.html">http://www.dnalc.org/view/16430-Gallery-19-DNA-model-1953.html</a>          2012/02/21 visited</p>	P21
P22. "Smithies uses starch-gel electrophoresis to identify protein polymorphisms."		<p>John C. Avise          1994. Molecular Markers, Natural History and Evolution, p. 42. Chapman &amp; Hall. New York, USA.  <a href="http://books.google.com.tw/books?id=aWSbZf-dRbEC&amp;printsec=frontcover&amp;hl=zh-TW#v=onepage&amp;q&amp;f=false">http://books.google.com.tw/books?id=aWSbZf-dRbEC&amp;printsec=frontcover&amp;hl=zh-TW#v=onepage&amp;q&amp;f=false</a></p> <p><i>It is used subject to the fair use doctrine of:</i>          •Taiwan Copyright Act Articles 52 &amp; 65</p>	P22
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P23. "Margoliash determines amino acid sequences for cytochrome c in several taxa and generates the first phylogenetic tree for a specific gene product."		<p>John C. Avise          1994. Molecular Markers, Natural History and Evolution, p. 42. Chapman &amp; Hall. New York, USA.  <a href="http://books.google.com.tw/books?id=aWSbZf-dRbEC&amp;printsec=frontcover&amp;hl=zh-TW#v=onepage&amp;q&amp;f=false">http://books.google.com.tw/books?id=aWSbZf-dRbEC&amp;printsec=frontcover&amp;hl=zh-TW#v=onepage&amp;q&amp;f=false</a></p> <p><i>It is used subject to the fair use doctrine of:</i>          •Taiwan Copyright Act Articles 52 &amp; 65</p>	P23
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		National Taiwan University Chau-Ti Ting	P25
		Walter M. Fitch and Emanuel Margoliash. 1967. Construction of phylogenetic trees. Science Vol. 155: p.282 <a href="http://www.sciencemag.org/content/155/3760/279.extract?sid=fe0a324a-4d85-453e-b77a-8a69782df245">http://www.sciencemag.org/content/155/3760/279.extract?sid=fe0a324a-4d85-453e-b77a-8a69782df245</a> 2012/02/21 visited <i>It is used subject to the fair use doctrine of:</i> •Taiwan Copyright Act Articles 52 & 65 <a href="http://www.sciencemag.org/content/155/3760/279.extract?sid=fe0a324a-4d85-453e-b77a-8a69782df245">Science /AAAS Copyright Statement</a>	P26
		Walter M. Fitch 1988. Citation Classic. No. 27. <a href="http://garfield.library.upenn.edu/classics.html">http://garfield.library.upenn.edu/classics.html</a> <a href="http://garfield.library.upenn.edu/classics1988/A1988N888200001.pdf">http://garfield.library.upenn.edu/classics1988/A1988N888200001.pdf</a> 2012/02/21 visited <i>It is used subject to the fair use doctrine of:</i> •Taiwan Copyright Act Articles 52 & 65 •The "Code of Best Practices in Fair Use for OpenCourseWare 2009 (" <a href="http://www.centerforsocialmedia.org/sites/default/files/10-305-OCW-Oc t29.pdf">http://www.centerforsocialmedia.org/sites/default/files/10-305-OCW-Oc t29.pdf</a> )" by A Committee of Practitioners of OpenCourseWare in the U.S. The contents are based on Section 107 of the 1976 U.S. Copyright Act	P27

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P28. "Kimura proposes that the majority of molecular changes in evolution are due to random drift of neutral or nearly neutral mutations."		<p>John C. Avise      1994. Molecular Markers, Natural History and Evolution, p. 42. Chapman &amp; Hall. New York, USA.  <a href="http://books.google.com.tw/books?id=aWSbZf-dRbEC&amp;printsec=frontcover&amp;hl=zh-TW#v=onepage&amp;q&amp;f=false">http://books.google.com.tw/books?id=aWSbZf-dRbEC&amp;printsec=frontcover&amp;hl=zh-TW#v=onepage&amp;q&amp;f=false</a></p> <p>2012/02/21 visited  <i>It is used subject to the fair use doctrine of:</i>      •Taiwan Copyright Act Articles 52 &amp; 65</p>	P28
P29. "Marshall Nirenberg, a young biochemist at the National Institute of Arthritic and Metabolic Diseases... Subsequently, within five years, the entire genetic code was deciphered"		<p>Genome News Network  <a href="http://www.genomenewsnetwork.org/resources/timeline/1961_Nirenberg.php">http://www.genomenewsnetwork.org/resources/timeline/1961_Nirenberg.php</a></p> <p>2012/02/21 visited  <i>It is used subject to the fair use doctrine of:</i>      •Taiwan Copyright Act Articles 52 &amp; 65</p>	P29
		<p>Cold Spring Harbor Laboratory  <a href="http://www.dnalc.org/view/16497-Gallery-22-Marshall-Nirenberg-1960.htm">http://www.dnalc.org/view/16497-Gallery-22-Marshall-Nirenberg-1960.htm</a></p> <p>2012/02/21 visited</p>	P29
		<p>Cold Spring Harbor Laboratory  <a href="http://www.dnalc.org/view/16498-Gallery-22-Marshall-Nirenberg-1999.htm">http://www.dnalc.org/view/16498-Gallery-22-Marshall-Nirenberg-1999.htm</a></p> <p>2012/02/21 visited</p>	P29
P30. "Nei's genetic distance has greatly facilitated the study of evolutionary relationships among populations or closely related species."		<p>M. Prakash      2007. Encyclopaedia of Gene Evolution Vol. 2, Molecular Genetics, p. 47. Discovery Publishing House. India.  <a href="http://books.google.com.tw/books?id=OoG4_y2Q9AEC&amp;printsec=frontcover&amp;hl=zh-TW#v=onepage&amp;q&amp;f=false">http://books.google.com.tw/books?id=OoG4_y2Q9AEC&amp;printsec=frontcover&amp;hl=zh-TW#v=onepage&amp;q&amp;f=false</a></p> <p>2012/02/21 visited  <i>It is used subject to the fair use doctrine of:</i>      •Taiwan Copyright Act Articles 52 &amp; 65</p>	P30
P31. "Molecular biologists by the 1970s had deciphered... developed new techniques for rapid DNA sequencing."		<p>Genome News Network  <a href="http://www.genomenewsnetwork.org/resources/timeline/1977_Gilbert.php">http://www.genomenewsnetwork.org/resources/timeline/1977_Gilbert.php</a></p> <p>2012/02/21 visited  <i>It is used subject to the fair use doctrine of:</i>      •Taiwan Copyright Act Articles 52 &amp; 65</p>	P31 38

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P32. “Palmer and colleagues begin an important series papers utilizing cpDNA for phylogenetic reconstructions in plants”		John C. Avise 1994. Molecular Markers, Natural History and Evolution, p. 42. Chapman & Hall. New York, USA. <a href="http://books.google.com.tw/books?id=aWSbZf-dRbEC&amp;printsec=frontcover&amp;hl=zh-TW#v=onepage&amp;q&amp;f=false">http://books.google.com.tw/books?id=aWSbZf-dRbEC&amp;printsec=frontcover&amp;hl=zh-TW#v=onepage&amp;q&amp;f=false</a> 2012/02/21 visited <i>It is used subject to the fair use doctrine of:</i> •Taiwan Copyright Act Articles 52 & 65	P32
			39