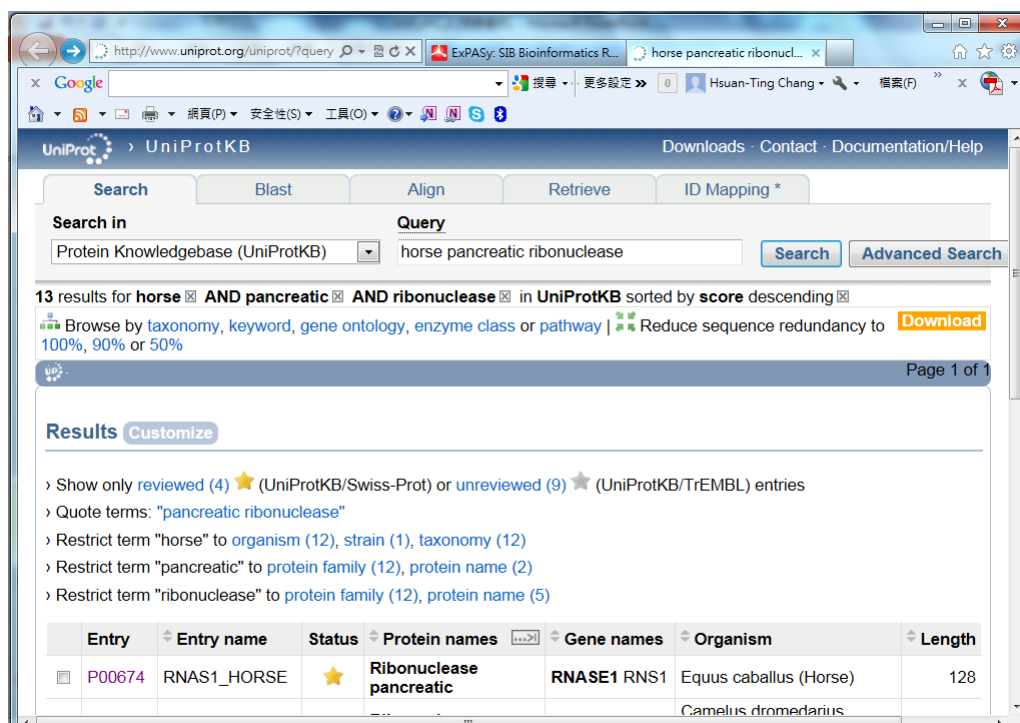
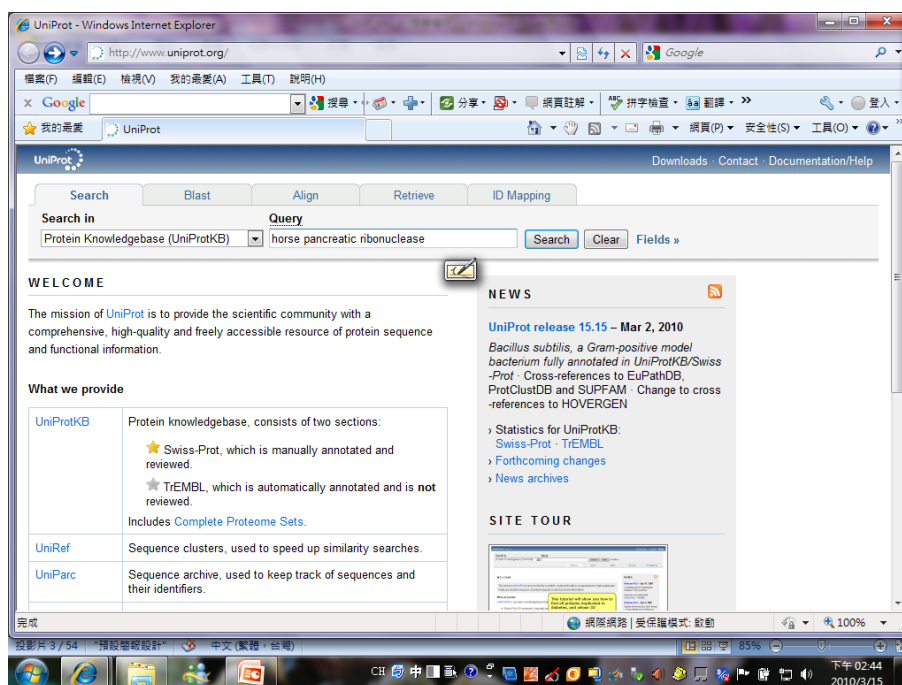


# Use of sequences to determine phylogenetic relationships

- Example 1.1 Retrieve the amino acid sequence of horse pancreatic ribonuclease
- ExPASy server: <http://www.expasy.org/>
- NCBI server: <http://www.ncbi.nlm.nih.gov/>
- Type the keywords: horse pancreatic ribonuclease (胰臟，核糖核酸酶)
- Select RNAS1\_HORSE and then find FASTA format





Ribonuclease pancreatic - Equus caballus (Horse) - Microsoft Internet Explorer

http://www.uniprot.org/uniprot/P00674

UniProtKB

Search in: Protein Knowledgebase (UniProtKB) Query: Search Clear Fields »

★ Reviewed, UniProtKB/Swiss-Prot P00674 (RNASE1\_HORSE)

Last modified March 3, 2009. Version 60. History...

Clusters with 100%, 90%, 50% identity | Documents (1) | Third-party data | Customize display

Names and origin

Protein names	Recommended name: <b>Ribonuclease pancreatic</b> EC=3.1.27.5 Alternative name(s): RNase 1 RNase A
Gene names	Name: <b>RNASE1</b> Synonyms: RNS1
Organism	<b>Equus caballus (Horse)</b>

Ribonuclease pancreatic - Equus caballus (Horse) - Microsoft Internet Explorer

http://www.uniprot.org/uniprot/P00674

Sequences

Sequence	Length	Mass (Da)	Tools
<input type="checkbox"/> P00674-1 [UniParc]	128	14,374	Blast

Last modified August 13, 1987. Version 1  
Checksum: A06727414097C1DD

```

10      20      30      40      50      60
KESPAMKFER QHMDSGSTSS SNPTTCNQMM KRRNMTGGWC KPVNTFVHEP LADVQAICLQ
70      80      90     100     110     120
KNITCKNGQS NCYQSSSSMH ITDCRLTSGS KYPNCAVQTS QKERHIIVAC EGNPTVPVHF
DASVEVST

```

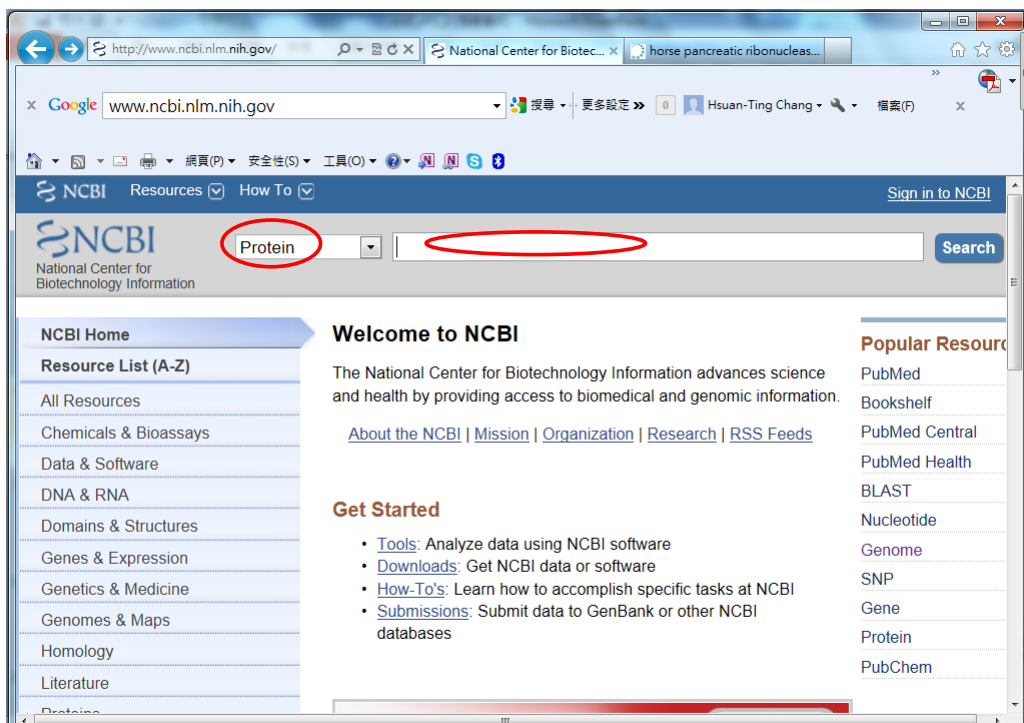
References

[1] "Horse pancreatic ribonuclease."  
Scheffer A.J., Beintema J.J.

## FASTA format

>sp|P00674|RNAS1\_HORSE Ribonuclease pancreatic (EC 3.1.27.5)  
(RNase 1) (RNase A) - Equus caballus (Horse).

```
KESPAMKFERQHMDSGSTSSSNPTYCNQMMKRRNMTQGWCK
PVNTFVHEPLADVQAICLQKNITCKNGQSNCYQSSSSMHITD
CRLTSGSKYPNCAYQTSQKERHIIVACEGNPYVPVHFDASVE
VST
```



http://www.ncbi.nlm.nih.gov/protein/?term=pancreatic+ribonuclease

Google P00674

Results: 18 Selected: 1

- ☐ [pancreatic ribonuclease \[Escherichia coli 3.2608\]](#)
  - 1. 268 aa protein
  - Accession: EIH56692.1 GI: 386179213
  - [GenPept](#) [FASTA](#) [Graphics](#) [Related Sequences](#) [Identical Proteins](#)
- ☐ [putative pancreatic ribonuclease \[Equus caballus\]](#)
  - 2. 165 aa protein
  - Accession: CAE45264.1 GI: 33945723
  - [GenPept](#) [FASTA](#) [Graphics](#) [Related Sequences](#) [Identical Proteins](#)
- ☒ [RecName: Full=Ribonuclease pancreatic; AltName: Full=RNase 1; AltName: Full=RNase A](#)
  - 3. 128 aa protein
  - Accession: P00674.1 GI: 133215
  - [GenPept](#) [FASTA](#) [Graphics](#) [Related Sequences](#)
- ☐ [angiogenin precursor \[Equus caballus\]](#)
  - 4. 146 aa protein
  - Accession: AAS15049.1 GI: 42411028

All (18)

- [Bacteria \(1\)](#)
- [Related Structures \(16\)](#)
- [RefSeq \(11\)](#)
- [Manage Filters](#)

Top Organisms [Tree]

- [Equus caballus \(16\)](#)
- [Camelus dromedarius \(1\)](#)
- [Escherichia coli 3.2608 \(1\)](#)

Analyze these sequences

- [Run BLAST](#)
- [Align sequences with COBALT](#)
- [Identify Conserved Domains with CD-Search](#)

http://www.ncbi.nlm.nih.gov/protein/P00674

Google www.ncbi.nlm.nih.gov

NCBI Resources How To

Protein Protein Limits Advanced

Display Settings: ☒ GenPept Send to: ☒ Change

**RecName: Full=Ribonuclease pancreatic; AltName: Full=RNase 1; AltName: Full=RNase A**

UniProtKB/Swiss-Prot: P00674.1

[FASTA](#) [Graphics](#)

Go to: ☒

LOCUS RNAS1\_HORSE 128 aa linear MAM 03-OCT-2012

DEFINITION RecName: Full=Ribonuclease pancreatic; AltName: Full=RNase 1; AltName: Full=RNase A.

ACCESSION P00674

VERSION P00674.1 GI:133215

DBSOURCE UniProtKB: locus RNAS1\_HORSE, accession P00674; class: standard.

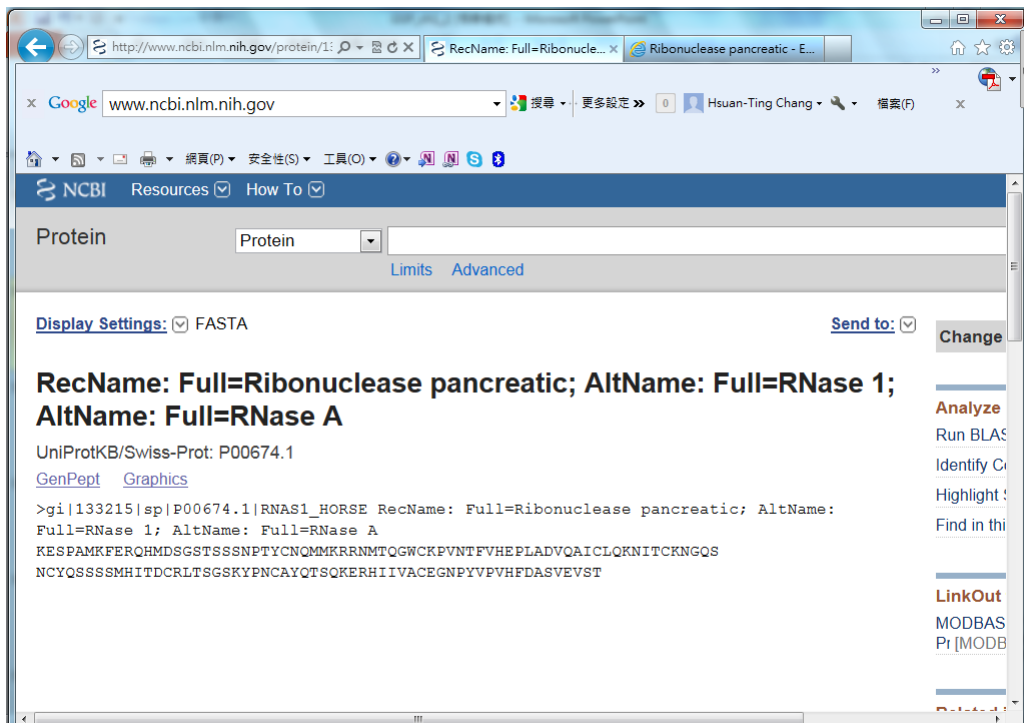
Created: Jul 21, 1986

Analyze

- [Run BLAST](#)
- [Identify Conserved Domains with CD-Search](#)
- [Highlight Conserved Domains](#)
- [Find in this database](#)

LinkOut

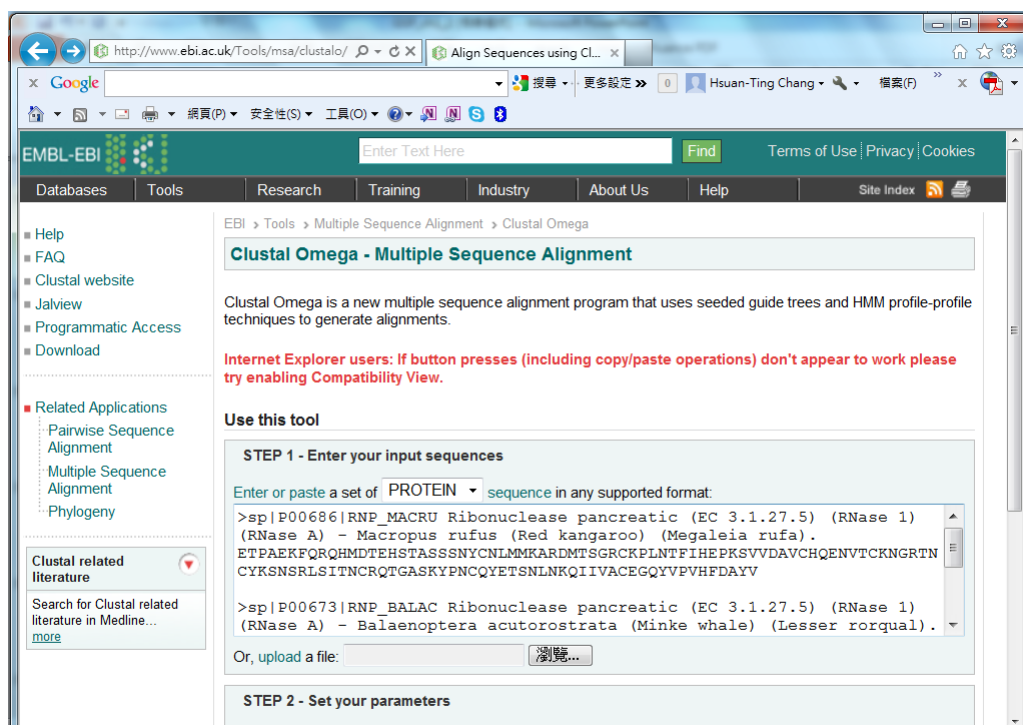
- [MODBAS](#)
- [PRIMODS](#)



## Example 1.2

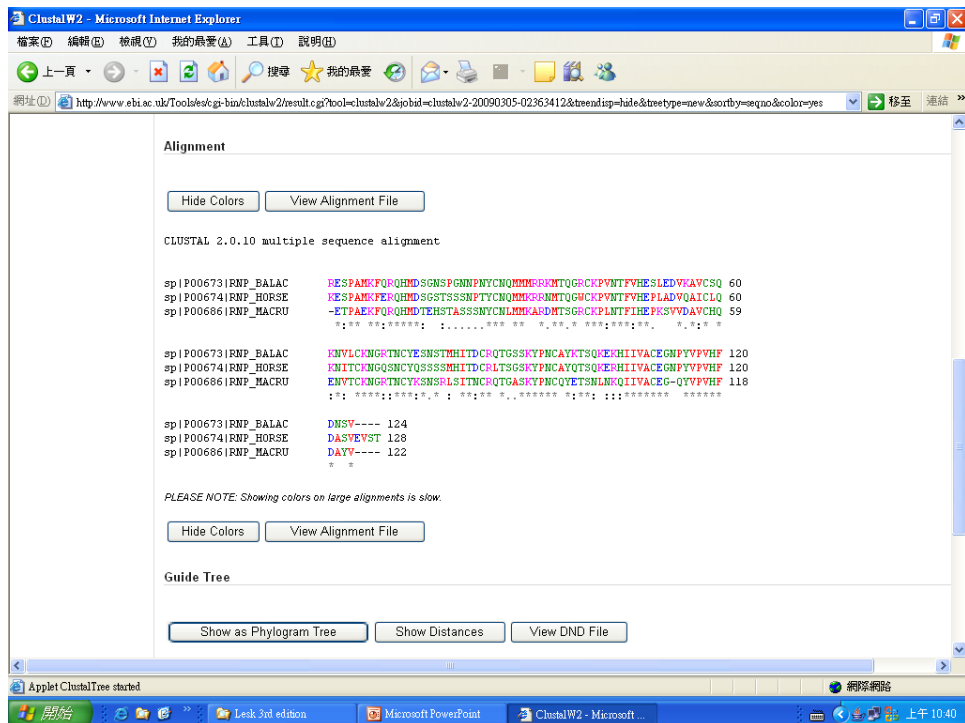
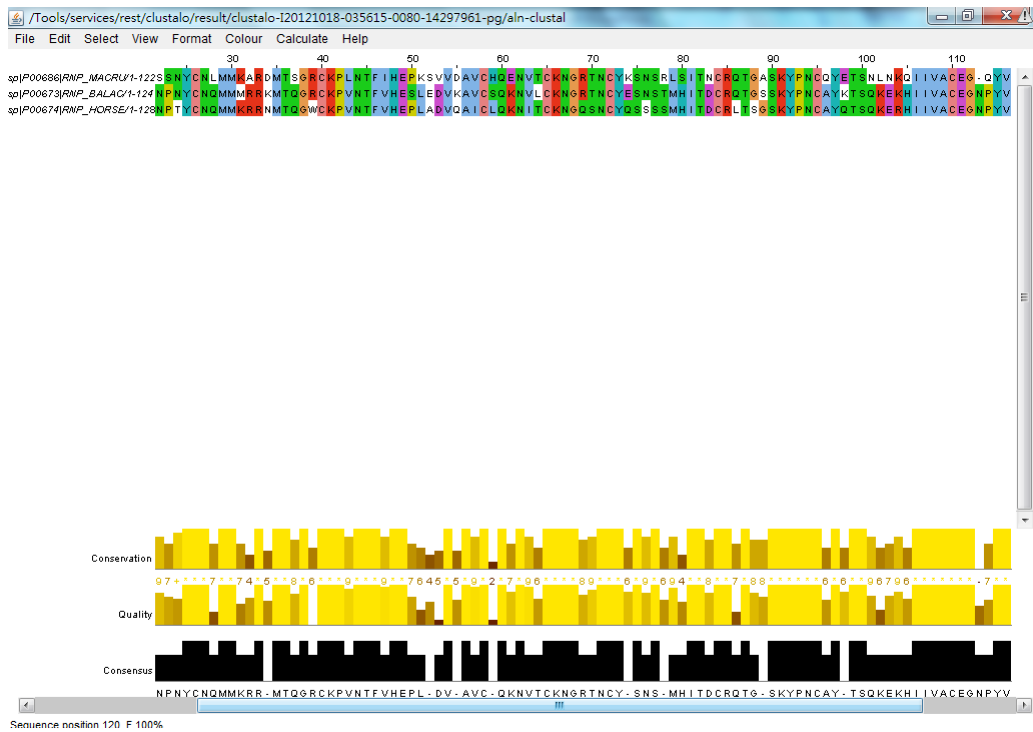
- Determine, from the sequences of pancreatic ribonuclease from *horse*, *minke whale* (小鬚鯨) and *red kangaroo*, which two of these species are most closely related.
- Retrieve the amino acid sequences from the database and use the **multiple sequence alignment** (MSA) tool **CLUSTAL-Omega**
  - <http://www.ebi.ac.uk/Tools/msa/>
  - New MSA tool that uses seeded guide trees and HMM profile-profile techniques to generate alignments. Suitable for medium-large alignments
- Clustal Omega is a multiple sequence alignment program for proteins. It produces biologically meaningful multiple sequence alignments of divergent sequences. evolutionary relationships can be seen via viewing Cladograms or Phylograms.

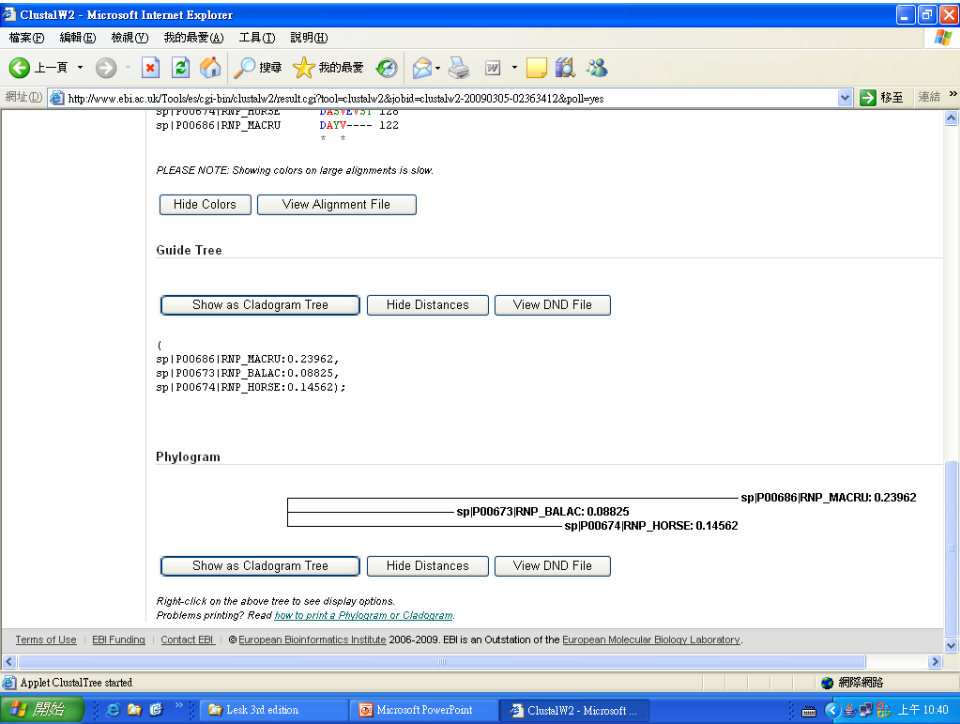
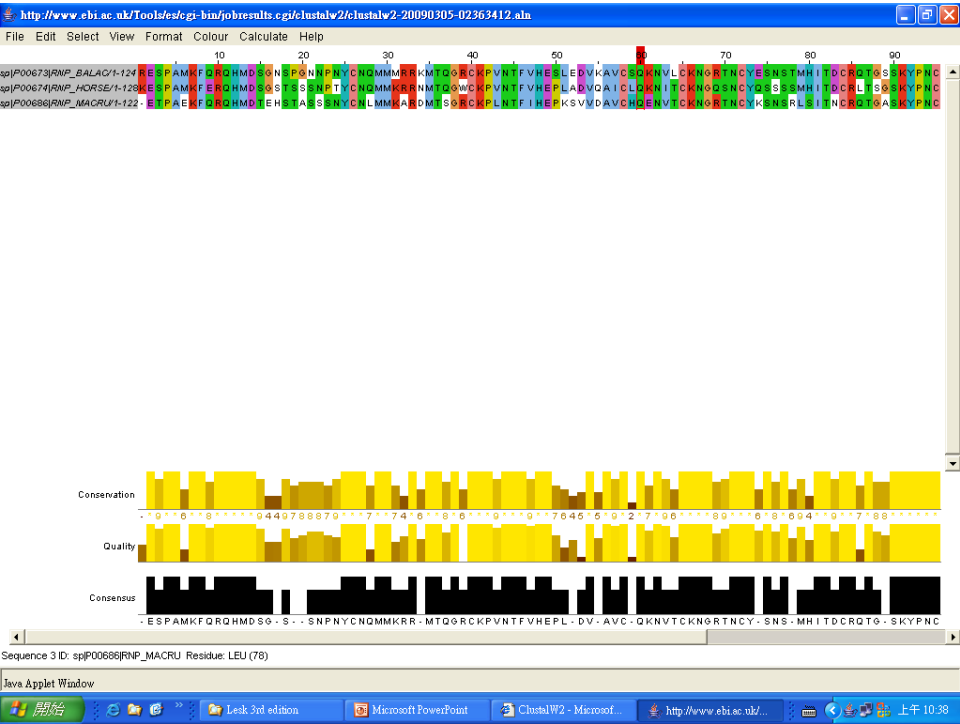
- >sp|P00686|RNP\_MACRU **Ribonuclease pancreatic** (EC 3.1.27.5) (RNase 1) (RNase A)  
- Macropus rufus (**Red kangaroo**) (Megaleia rufa).
- ETPAEKFQRQHMDTEHSTASSSNYCNLMMKARDMTSGRCKPLNTFIHEPKSVV  
DAVCHQENVCTCKNGRTNCYKSNRSLITNCRQTGASKYPNCQYETSNLNKQIIV  
ACEGQYVPVHFDAYV
- >sp|P00673|RNP\_BALAC **Ribonuclease pancreatic** (EC 3.1.27.5) (RNase 1) (RNase A)  
- Balaenoptera acutorostrata (**Minke whale**) (Lesser rorqual).
- RESPAMKFQRQHMDSGNSPGNNPNYCNQMMMRRKMTQGRCKPVNTFVHESL  
EDVKAVCSQKNVLCKNGRTNCYESNSTMHITDCRQTGSSKYPNCAYKTSQKEK  
HIIVACEGNPYVPVHFDNSV
- >sp|P00674|RNP\_HORSE **Ribonuclease pancreatic** (EC 3.1.27.5) (RNase 1) (RNase A)  
- Equus caballus (**Horse**).
- KESPAMKFERQHMDSGSTSSSNPTYCNQMMKRRNMTQGWCKPVNTFVHEPLA  
DVQAICLQKNITCKNGQSNCYQSSSMHITDCRLTSGSKYPNCAYQTSQKERHII  
VACEGNPYVPVHFDASVEVST











## Alignment Result

- Horse and whale share the most identical residues. The result appears significant, and therefore confirm our expectation.
  - Knowing that horse and whale are placental (有胎盤的) mammals and kangaroo is a marsupial (有袋動物), we expect horse and whale to be closer pair.

## FASTA format description

A sequence in FASTA format begins with a single-line description, followed by lines of sequence data. The description line is distinguished from the sequence data by a greater-than (">") symbol in the first column. It is recommended that all lines of text be shorter than 80 characters in length. An example sequence in FASTA format is:

```
>gi|532319|pir|TVFV2E|TVFV2E envelope protein
ELRLRYCAPAGFALLKCNDADYDGFKTNCNSVSVVHCTNLMNTTVTTGLLNGSYSENRT
QIWQKHRTSNDALILLNKHYNLTVCKRPGNKTVLPVTIMAGLVFHSQKYNLRLRQAWC
HFPSNWKGAWEVKEEIVNLPKERYRGTNDPKRIFFQRQWGDPEANLWFNCHGEFFYCK
MDWFLNYLNNLTVDADHNECKNTSGTKSGNKRAPGPCVQRTYVACHIRSVIWLLETISKK
TYAPPREGHLECTSTVTGMTVELNYIPKNRTNVTLSQIESIWAELDRYKLVEITPIGF
APTEVRRYTGGERQKRVPFVXXXXXXXXXXXXXXXXXXXXXXXXXVQSQHLLAGILQQQKNL
LAAVEAQQQMLKLTWGVK
```

In bioinformatics, a **sequence alignment** is a way of arranging the sequences of DNA, RNA, or protein to identify regions of similarity that may be a consequence of functional, structural, or evolutionary relationships between the sequences. Aligned sequences of nucleotide or amino acid residues are typically represented as rows within a matrix. Gaps are inserted between the residues so that identical or similar characters are aligned in successive columns.

### Global and local alignments

```
Global  FTFTALILLAVAV
        F--TAL-LLA-AV

Local   FTFTALILL-AVAV
        --FTAL-LLAAV--
```

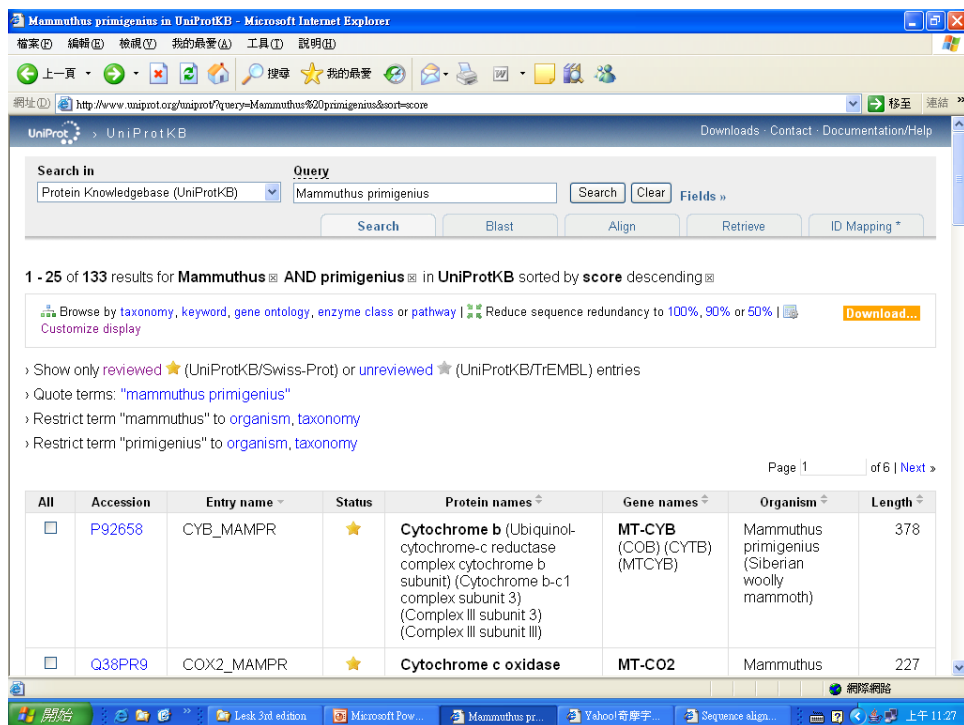
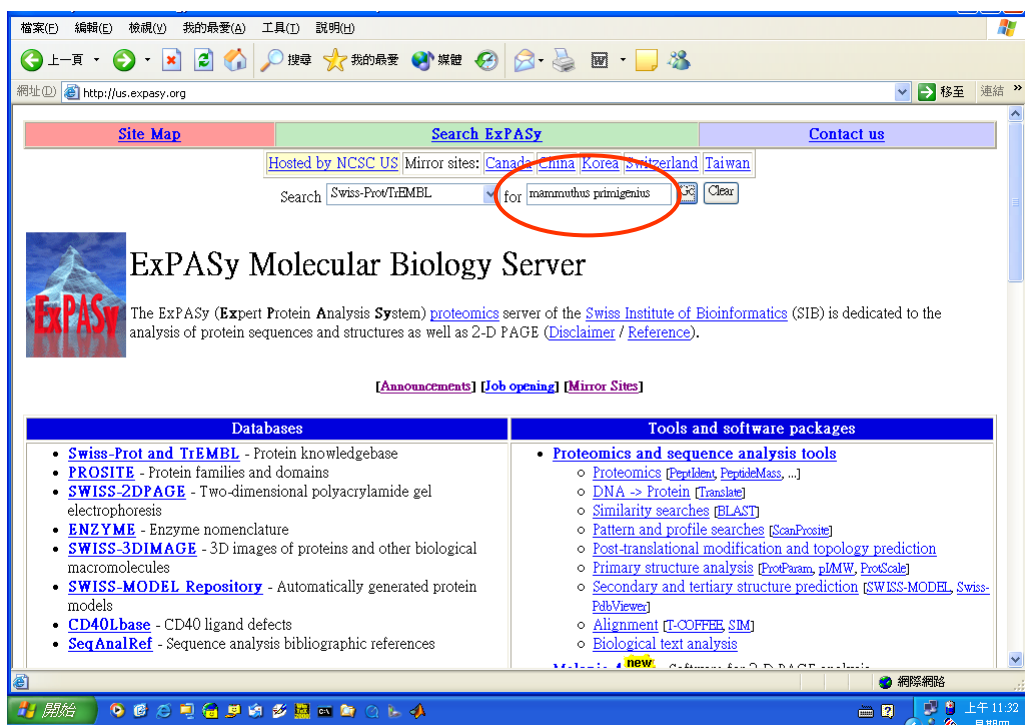
## Homework #1

- Find the pancreatic ribonuclease of any other three species and perform following operations:
- 1. Perform multiple sequence alignment by use of CLUSTAL-W software
- 2. Find the corresponding DNA sequences.
- 3. Perform multiple sequence alignment for retrieved DNA sequences.

## Case Study 1.5 – A harder one

- Two living genera of elephant: the African elephant and the Indian
- 藉由北極長毛象(*Mammuthus primigenius*)化石之粒腺體細胞色素b (mitochondrial cytochrome b)，界定出哪一種現代大象(*Loxodonta africana* and *Elephas maximus*)與遠古之長毛象有較近之血緣關係。
- From the results, it appears that mammoth is more closely related to African elephant. However, there are few differences. Are they significant?
- Questions:
  - Could we tell from these sequences alone that they are from closely related species?
  - Given the differences are small, do they represent evolutionary divergence arising from selection, or merely random noise or drift?

- >sp|P92658|CYB\_MAMPR Cytochrome b OS=Mammuthus primigenius GN=MT-CYB PE=3 SV=3
- MTHIRKSHPLLKILNKSFIDLTPSNISTWWNFGSLLGACLITQILTGLFLAMHYTPDTM  
TAFSSMSHICRDVNYGWIRQLHSNGASIFFLCYTHIGRNIYYGSYLYSETWNTGIMLL  
LITMATAFMGYVLPWGQMSFWGATVITNLFSAIPYIGTDLVEWIWGGFSVDKATLNRFFA  
LHFILPFTMIALAGVHLTFLHETGSNNPLGLTSDSDKIPFHPYYTIKDFLGLLILILFLL  
LLALLSPDMLGDPDNYPADPLNTPLHIKPEWYFLFAYAILRSVPNKLGGVLALLLSILI  
LGIPLLHTSKHRSMMLRPLSQVLFWTLATDLLMLTWIGSQPVEYPYIIIGQMASILYFS IILAFLPIAGMIENYLIK
- >sp|P24958|CYB\_LOXAF Cytochrome b OS=Loxodonta africana GN=MT-CYB PE=3 SV=2
- MTHIRKSHPLLKIINKSFIDLTPSNISTWWNFGSLLGACLITQILTGLFLAMHYTPDTM  
TAFSSMSHICRDVNYGWIRQLHSNGASIFFLCYTHIGRNIYYGSYLYSETWNTGIMLL  
LITMATAFMGYVLPWGQMSFWGATVITNLFSAIPYIGTNLVEWIWGGFSVDKATLNRFFA  
LHFILPFTMIALAGVHLTFLHETGSNNPLGLTSDSDKIPFHPYYTIKDFLGLLILILLLL  
LLALLSPDMLGDPDNYPADPLNTPLHIKPEWYFLFAYAILRSVPNKLGGVLALLLSILI  
LGLMPLLHTSKHRSMMLRPLSQVLFWTLTMDLLTLTWIGSQPVEYPYIIIGQMASILYFS IILAFLPIAGVIENYLIK
- >sp|O47885|CYB\_ELEMA Cytochrome b OS=Elephas maximus GN=MT-CYB PE=3 SV=1
- MTHTRKFHPLFKIINKSFIDLTPSNISTWWNFGSLLGACLITQILTGLFLAMHYTPDTM  
TAFSSMSHICRDVNYGWIRQLHSNGASIFFLCYTHIGRNIYYGSYLYSETWNTGIMLL  
LITMATAFMGYVLPWGQMSFWGATVITNLFSAIPYIGTNLVEWIWGGFSVDKATLNRFFA  
LHFILPFTMVALAGVHLTFLHETGSNNPLGLTSDSDKIPFHPYYTIKDFLGLLILILLLL  
LLALLSPDMLGDPDNYPADPLNTPLHIKPEWYFLFAYAILRSVPNKLGGVLALFLSILI  
LGLMPLLHTSKHRSMMLRPLSQVLFWTLTMDLLTLTWIGSQPVEHPYIIIGQMASILYFS IILAFLPIAGMIENYLIK



Cytochrome b - Mammothus primigenius (Siberian woolly mammoth) - Microsoft Internet Explorer

檔案(F) 編輯(E) 檢視(V) 我的最愛(A) 工具(T) 說明(H)

地址(1) http://www.uniprot.org/uniprot/P92658

★ Reviewed, UniProtKB/Swiss-Prot **P92658** (CYB\_MAMPR)

Last modified March 3, 2009. Version 56. [History...](#)

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Clusters with 100%, 90%, 50% identity | Documents (1) | Third-party data | Customize display

[TEXT](#) [XML](#) [RDF/XML](#) [GFF](#) [FASTA](#)

[Names and origin](#) · [Protein attributes](#) · [General annotation \(Comments\)](#) · [Ontologies](#) · [Sequence annotation \(Features\)](#) · [Sequences](#) · [References](#) · [Cross-references](#) · [Entry information](#) · [Relevant documents](#)

**Names and origin** Hide | Top

Protein names	<p>Recommended name:  <b>Cytochrome b</b></p> <p>Alternative name(s):  Ubiquinol-cytochrome-c reductase complex cytochrome b subunit  Cytochrome b-c1 complex subunit 3  Complex III subunit 3  Complex III subunit III</p>
Gene names	<p>Name: <b>MT-CYB</b></p> <p>Synonyms: COB, CYTB, MTCYB</p>
Encoded on	Mitochondrion
Organism	<b>Mammothus primigenius (Siberian woolly mammoth)</b>
Taxonomic identifier	37349 [NCBI]
Taxonomic lineage	Eukaryota > Metazoa > Chordata > Craniata > Vertebrata > Euteleostomi > Mammalia > Eutheria > Afrotheria > Proboscidea > Elephantidae > Mammuthus

開始 樂高 3rd edition Microsoft Pow... Cytochrome b... Yahoo!奇摩字... Sequence align... 上午 11:28

檔案(F) 編輯(E) 檢視(V) 我的最愛(A) 工具(T) 說明(H)

地址(1) http://www.ebi.ac.uk/clustalw/

def	def	def	def	def
-----	-----	-----	-----	-----

TREE GRAPH		PHYLOGENETIC TREE		
TYPE	DISTANCES	TREE TYPE	CORRECT DIST.	IGNORE GAPS
cladogram	hide	none	off	off

Enter or Paste a set of Sequences in any supported format: [Help](#)

```
>sp|O47865|CYB_ELEMA Cytochrome b - Elephas
maximus (Indian elephant).
MTHTKRFHPLFKIINKSFIDLPTSNISTWWNFSGLLGACLTQIL
TGLFLAMHYTPDTMTAFSSMHSICRDVNYGWIIRQLHNSGASIFFL
CLYTHIGRNIYYGSYLYSETWNTGIMLLITMTAFHMGVLPUGQM
SFUGATVITNLFSAPYIGTNLVEWINGGFSVDKATLNRFPAHF I
LPFTMVALAGVHLTFLHETGSNNPLGLTSDSKIPFPYTYIKDFL
GLLILILLLLLLALLSPDMLGDPDNYMPADPLNTPLHIKPEWYFLF
AYAILRSVPNKLGGVLALFLSILILGLMPLLLHTSKHRSMLRPLSQ
VLFWTLTMDLLTLTWIGSQVHPHYIIIGQMASILYFSIILAFLEPI
```

Upload a file:  [瀏覽...](#) [Run](#) [Reset](#)

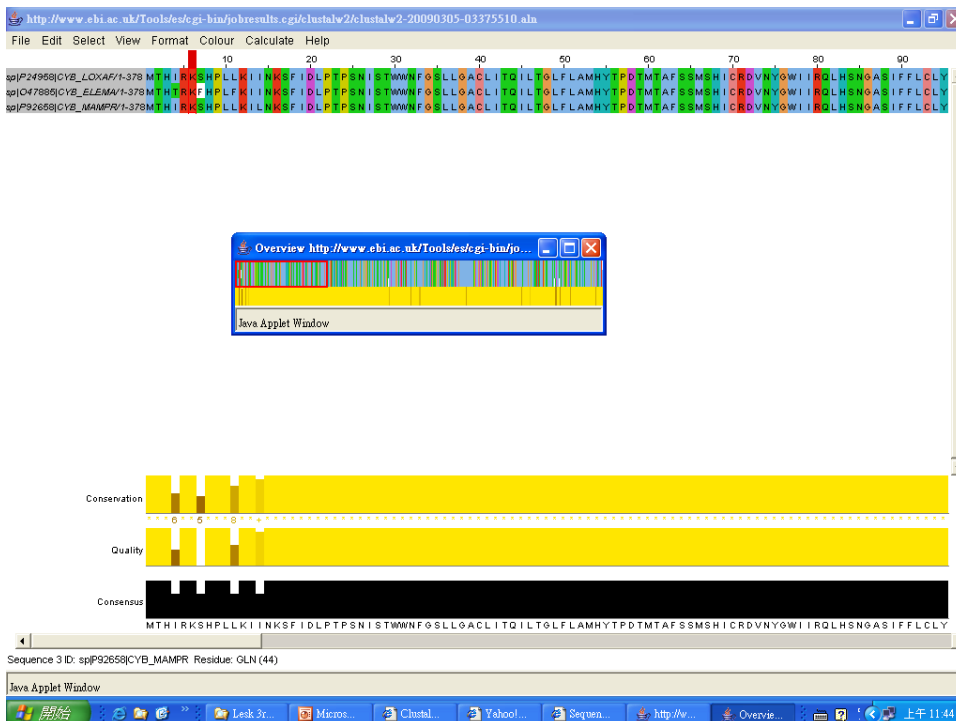
If you plan to use these services during a course please contact us using the email below.

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完成 網路網站 上午 11:22 星期四







## Background: Similarity & Homology

- **Similarity**: the observation or measurement of resemblance (相似點) and difference, independent of the source of the resemblance.
- **Homology**: the sequences and the organisms in which they occur are descended from a common ancestor, with the implication that the similarities are shared ancestral characteristics.
  - Homology must be an *inference* from observations of similarity. Only a few special cases is homology directly observables.
- The need for thoughtful scientific judgment!

- Does the very high degree of similarity of the sequences justify the conclusion that they are homologous; or are there other explanations?
- A functional cytochrome b *requires* so many conserved residues
- It starts out from independent ancestors and that common selective pressures forced them to become similar??
- Indian elephant has evolved faster than that of the African elephant or the mammoth, accumulating more mutations
- They gained a common gene by transfer from an unrelated organism via a virus.

## SINES & LINES

- Short and long interspersed nuclear elements – repetitive non-coding sequences that form large fractions of eukaryotic genomes. 30% of human chromosomal DNA, and over 50% of some higher plant genomes.
- Typically,
  - SINES are ~70 – 500 bps long, up to  $10^6$  copies may appear.
  - LINES may be up to 7000 bps long, up to  $10^5$  copies may appear.
- To derive the phylogenetic relationship
- Features of SINES that make them useful for phylogenetic studies include:
  - A SINE is either present or absent.

- SINES are inserted at random in the non-coding portion of a genome.
  - Appearance of similar SINES at the same locus in two species implies that the species share a common ancestor in which the insertion event occurred.
- SINE insertion appears to be irreversible: no mechanism for loss of SINES is known, other than rare large-scale deletions that include SINE.
- Not only do SINES shown relationships, they imply which species came first. The last common ancestor of species containing a common SINE must have come after the last common ancestor linking these species and another that lacks this SINE.

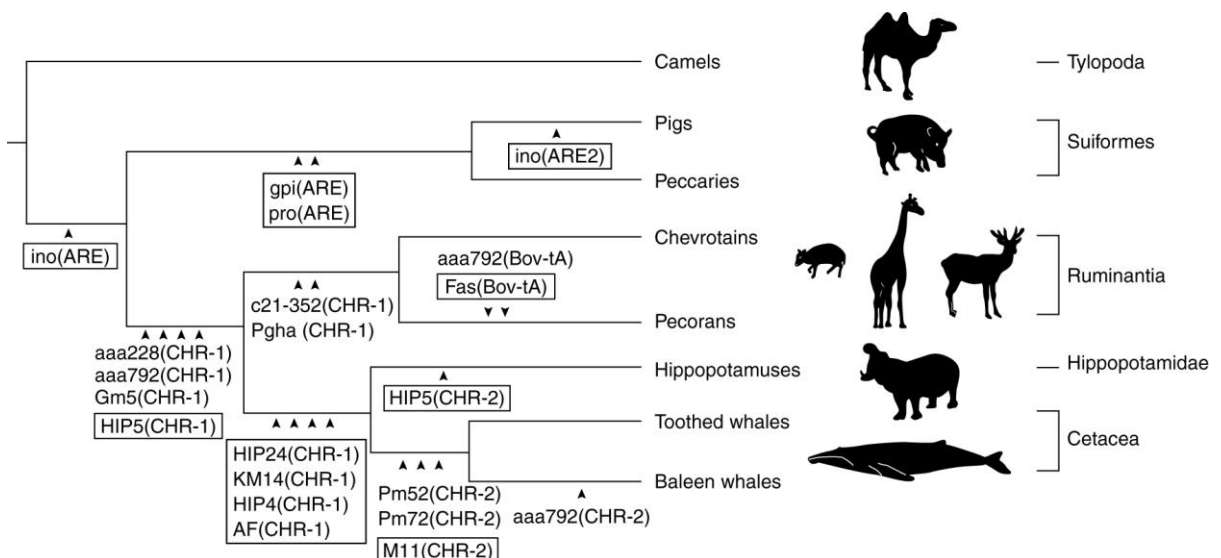
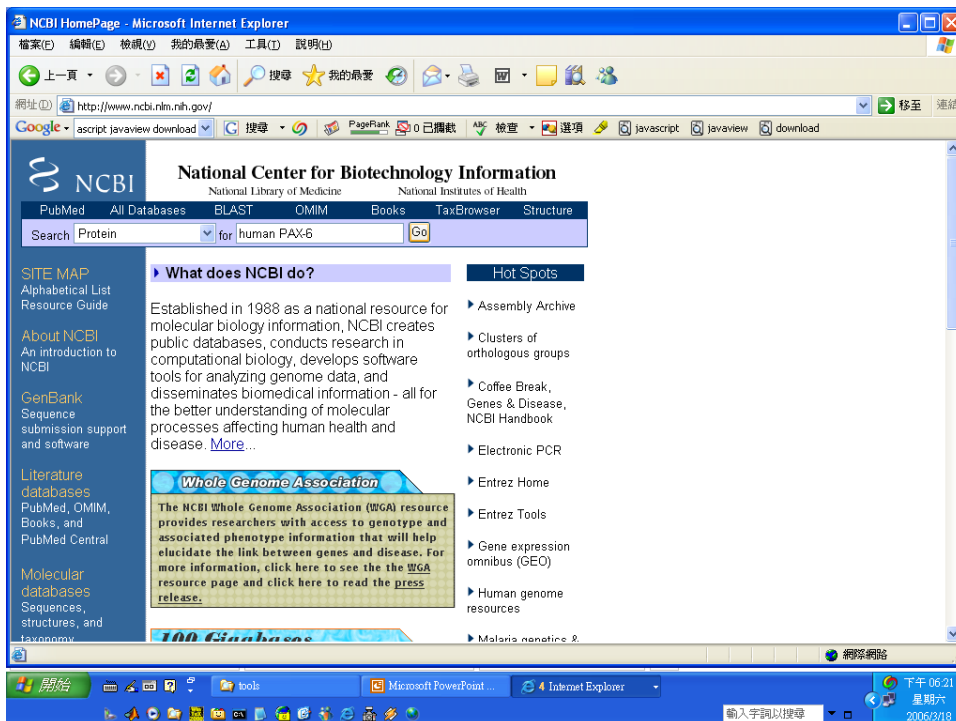


Fig. 1.5 Phylogenetic relationships among cetaceans and other artiodactyl subgroups, derived from analysis of SINE sequences.

# Searching for similar sequences in databases: PSI-BLAST

- The search of a database for items similar to a probe.
- The ideal method is both sensitive and selective:
  - Sensitive: it picks up even very distant relationships
  - Selective: all the relationships that it reports are true
- A powerful tool from NCBI USA: PSI-BLAST
  - Position Specific Iterated – basic linear alignment sequence tool
- Example 1.4. Homologues of the human PAX-6 gene.
- Example 1.5. What species contain homologues of human PAX-6 detectable by PSI-BLAST?



http://www.ncbi.nlm.nih.gov/protein/7t... human PAX-6 - Protein - ...

Results: 1 to 20 of 67

Chain A, Solution Structure Of The Homeobox Domain Of The Human Paired Box Protein Pax-6

1. Protein Pax-6  
80 aa protein  
Accession: 2CUE\_A GI: 159163961  
GenPept FASTA Graphics Related Sequences

Chain A, Crystal Structure Of The Human Pax-6 Paired Domain-Dna Complex Reveals A General Model For Pax Protein-Dna Interactions

2. 133 aa protein  
Accession: 6PAX\_A GI: 5822580  
GenPept FASTA Graphics Related Sequences

RecName: Full=Paired box protein Pax-6; AltName: Full=Aniridia type II protein; AltName: Full=Oculorhombin

3. 422 aa protein  
Accession: P26367.2 GI: 6174889  
GenPept FASTA Graphics Related Sequences Identical Proteins

RecName: Full=Paired box protein Pax-6; AltName: Full=Oculorhombin

4. 422 aa protein  
Accession: P63015.1 GI: 51702790  
GenPept FASTA Graphics Related Sequences Identical Proteins

Top Organisms [Tree]  
Homo sapiens (27)  
Mus musculus (14)  
Pediculus humanus corporis (6)  
Schistosoma mansoni (3)  
Chrysopa quinquecirrha (3)  
All other taxa (14)  
More...

Find related data  
Database: Select  
Find items

Search details  
("Homo sapiens" [Org]) NUM LOCK: ON

NCBI Sequence Viewer v2.0 - Microsoft Internet Explorer

Search Protein for [ ] Go Clear

Display FASTA Show 5 Send to [ ]

Range from begin to end Refresh

1: P26367 Reports Paired box protei..[6174889]

BLINK, Conserved Domains, Links

>gi|6174889|sp|P26367|PAX6\_HUMAN Paired box protein Pax-6 (Oculorhombin) (Aniridia type II protein)  
MQNSHSGVNLGGVFNGLPLDSTRQKIVELAHSGARPCDISRILQVSNCGCVSKILGRYYETGSIRFRA  
IGGSRFRVATPEVVSIAQYKRECPFIFAEIRDRLLSEGVCTNDNIPSVSSINRVLNNLASEKQMGAD  
QBYDLRLMLNGTSGWTRPGVYPTSPVQPTQDCCQCGGGGENTSSISNEDSDSAQRLQLKRL  
QNRNFTTQEQIALKLEKFERHTYPTDVFARELAALIDLEARIQVWFSNRRAKVRREKLNRORQASN  
TPSHIPSSSFSTSVYQPIPOPTTPVSSFTSGMLGRDITALTNTYSALPPNPSFTMANLPMOPPVPSQ  
TSSYSCHMLPTSPSVNGRSYDTTPPHMQTHMNSQPNGTSGTTSTGLISPGVSVVQVPGSEPDMSQYWR  
LQ

Disclaimer | Write to the Help Desk  
NCBI | NLM | NIH

Mar 14 2008 11:21:02

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http://www.ncbi.nlm.nih.gov/protein/61

RecName: Full=Paired box protein Pax-6; AltName: Full=Aniridia type II protein; AltName: Full=Oculorhombin

UniProtKB/Swiss-Prot: P26367.2

FASTA

Send to: Change registry

Analyze this  
Run BLAST  
Identify Conserved Domains  
Highlight Sequences  
Find in this Sequence

Protein 3D Structure

gi|6174889|sp|P26367.2|PAX6\_HUMAN RecName: Full=Paired box protein Pax-6; AltName: Full=Aniridia type II protein; AltName: Full=Oculorhombin

QNSHSGVNLGGVFVNGRPLDPSTRQKIVELAHSGARPCDISRIQVSNQCVSKILGRYYETGSIRPRA  
GGSKPRVATPEVVSKIAQYKRECPISFAWEIRDRLLESEGVCNDNDNIPSVSSINRVLNLAKEQQMGA  
MYDKLRMLNGQTGSWGRPGWYPGTSVPGQPTQDGCQQQEGGENTNSISSNGEDSDEAQMRLQKRKL  
RNRSTFTQEQIEALEKEFERTHYPDVFAERERLAAKIDLPEARIQVWFSNRRAKWRREEKLRNQRQAS  
PSHIPISSSFTSVYQPIQPPTTPVSSFTSGSMLGRITDALTNTYSALPPMPSFTMANNLPMOPFVPSQ  
SSYSCLMPTSPSVNGRSYDTYTPPHMQTHMNSQPMQ

BLAST: Basic Local Alignment Search Tool - Microsoft Internet Explorer

http://blast.ncbi.nlm.nih.gov/Blast.cgi

Basic BLAST

Choose a BLAST program to run.

nucleotide blast Search a nucleotide database using a nucleotide query  
Algorithms: blastn, megablast, discontinuous megablast

protein blast Search protein database using a protein query  
Algorithms: blastp, psi-blast, phi-blast

blastx Search protein database using a translated nucleotide query

tblastn Search translated nucleotide database using a protein query

tblastx Search translated nucleotide database using a translated nucleotide query

Specialized BLAST

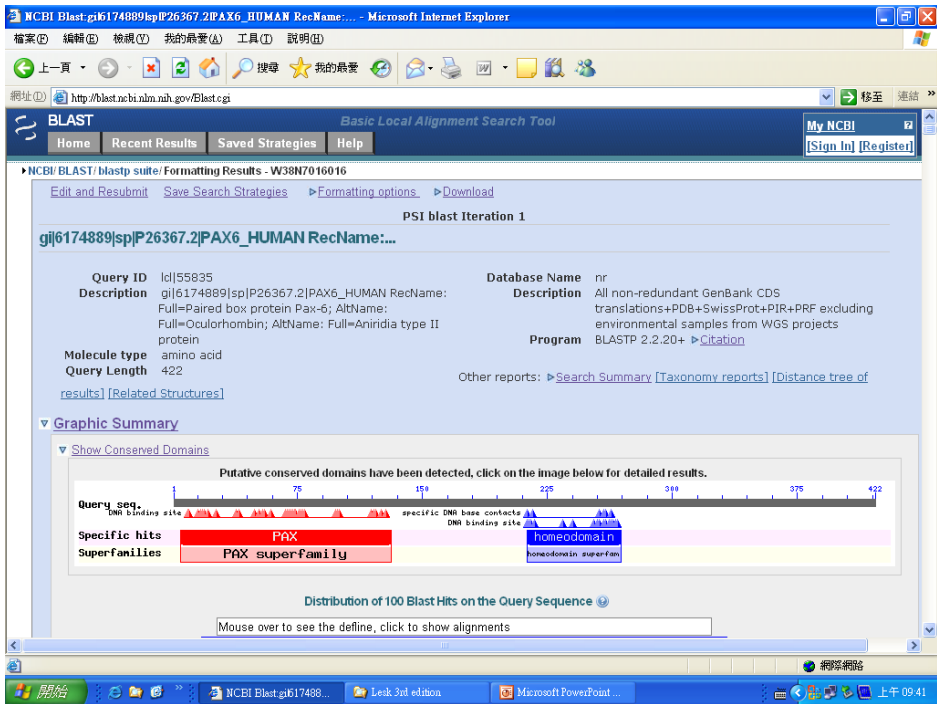
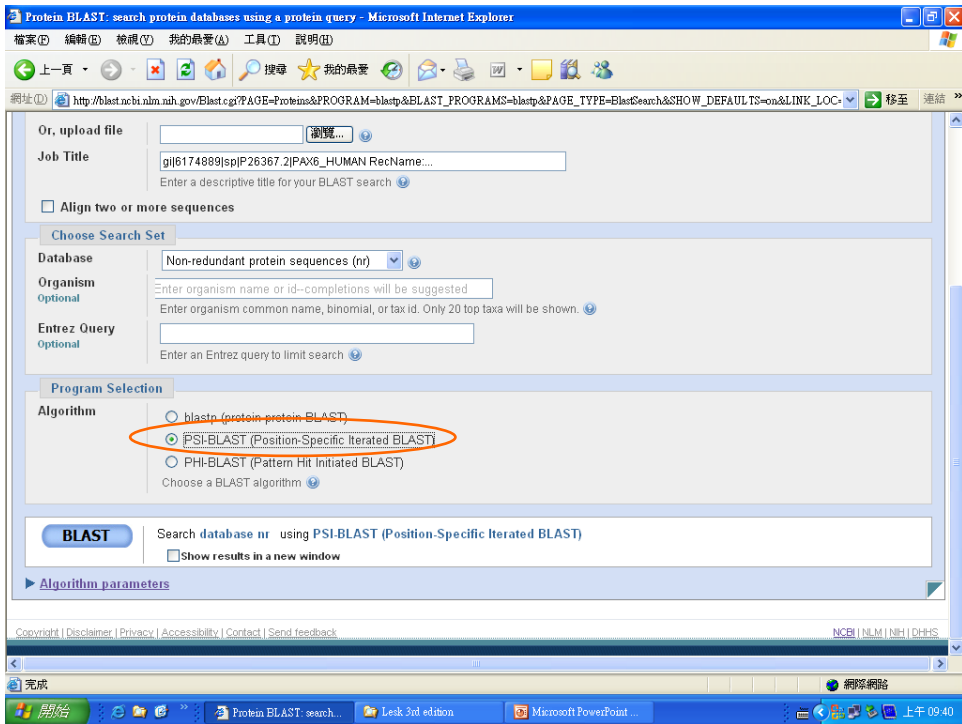
Choose a type of specialized search (or database name in parentheses.)

- Make specific primers with [Primer-BLAST](#)
- Search [trace archives](#)
- Find [conserved domains](#) in your sequence (cds)
- Find sequences with similar [conserved domain architecture](#) (cdart)
- Search sequences that have [gene expression profiles](#) (GEO)
- Search [immunoglobulins](#) (IgBLAST)
- Search for [SNPs](#) (snp)
- Screen sequence for [vector contamination](#) (vecscreen)
- [Align](#) two (or more) sequences using BLAST (bl2seq)
- Search [protein](#) or [nucleotide](#) targets in PubChem BioAssay

Relationships Between Sequences

The new Tree View option on the NCBI Web BLAST service presents a dendrogram or tree display that clusters sequences according to their distances from the query sequence.

[More tips...](#)



Sequences with E-value BETTER than threshold

Sequences producing significant alignments:

	Score (Bits)	E Value
<a href="#">ref NP_000271.1 </a> paired box gene 6 isoform a [Homo sapiens] >...	870	0.0
<a href="#">gb EAW68233.1 </a> paired box gene 6 (aniridia, keratitis), isofo...	869	0.0
<a href="#">ref NP_001035735.1 </a> paired box gene 6 [Bos taurus] >sp Q1L2F1...	868	0.0
<a href="#">gb AAA59962.1 </a> oculorhombin >gb AAA59963.1  oculorhombin	868	0.0
<a href="#">gb EDL79721.1 </a> paired box gene 6, isoform CRA_a [Rattus norve...	868	0.0
<a href="#">ref NP_037133.1 </a> paired box 6 [Rattus norvegicus] >sp P63016...	868	0.0
<a href="#">gb ABA90484.1 </a> paired box protein PAX6 isoform a [Oryctolagus...	867	0.0
<a href="#">dbj BA052023.1 </a> unnamed protein product [Homo sapiens]	867	0.0
<a href="#">dbj BAC25729.1 </a> unnamed protein product [Mus musculus]	867	0.0
<a href="#">gb ABI98848.1 </a> paired box 6 transcript variant 3 [Columba livia]	865	0.0
<a href="#">pfl 19023284</a> PAX6 gene	863	0.0
<a href="#">ref NP_001595.2 </a> paired box gene 6 isoform b [Homo sapiens] >...	863	0.0
<a href="#">ref NP_038655.1 </a> paired box gene 6 [Mus musculus] >emb CAA453...	862	0.0
<a href="#">ref NP_001091013.1 </a> paired box gene 6 [Canis lupus familiaris...	861	0.0
<a href="#">gb EDL27748.1 </a> paired box gene 6, isoform CRA_d [Mus musculus]	861	0.0
<a href="#">ref EAW68236.1 </a> paired box gene 6 (aniridia, keratitis), isofo...	861	0.0
<a href="#">gb EDL79723.1 </a> paired box gene 6, isoform CRA_c [Rattus norve...	860	0.0
<a href="#">emb CAC80516.1 </a> paired box protein [Mus musculus]	860	0.0
<a href="#">gb AAS48919.1 </a> paired box 6 isoform 5a [Rattus norvegicus] >g...	859	0.0
<a href="#">ref NP_001075686.1 </a> paired box protein PAX6 isoform b [Orycto...	858	0.0
<a href="#">emb CAE45868.1 </a> hypothetical protein [Homo sapiens]	858	0.0
<a href="#">gb ABS17534.1 </a> PAX6 [Bufo raddei]	858	0.0

CLUSTAL W (1.82) multiple sequence alignment

gi 2133658 pir  I45557	MFTLQPTPTAIGTVVPWSAGTLIERLPSELDMAHKGHSQVNLGGVFG	50
sp P26367 PAX6_HUMAN	-----MQNSHSQVNLGGVFN	17
	..*****	
gi 2133658 pir  I45557	GRPLPDSTRQKIVELAHSGARPCDISRILQVSNQCVSKILGRYYETGSIR	100
sp P26367 PAX6_HUMAN	GRPLPDSTRQKIVELAHSGARPCDISRILQVSNQCVSKILGRYYETGSIR	67
	*****	
gi 2133658 pir  I45557	PRAIGGSKPRVATAEVVSKISQYKRECPISFAWEIRDRLLENQVCTNDNI	150
sp P26367 PAX6_HUMAN	PRAIGGSKPRVATPEVVKIAQYKRECPISFAWEIRDRLLENQVCTNDNI	117
	*****	
gi 2133658 pir  I45557	PSVSSINRVLRLNLAQKEQQSTGSGSSSTSAGNSISAKVSVSIGNVSNV	200
sp P26367 PAX6_HUMAN	PSVSSINRVLRLNLAQKEQQ-----HG-----	138
	*****:;*	
gi 2133658 pir  I45557	ASGSRGTLSSSTDLMQTATPLNSESAGGATNSGEGSEQEAIYEKRLRLNT	250
sp P26367 PAX6_HUMAN	-----ADGHYDKLRMLNG	151
	..:;***:	
gi 2133658 pir  I45557	QHAAGPGLPEPARAAPLVGQSPNHLGTRSSHPQLVGHNHQALQQHQQSW	300
sp P26367 PAX6_HUMAN	QTGS-----WGTRP-----	160
	*.:***	
gi 2133658 pir  I45557	PPRHYSGSWYPTSLSEIPISAPNIAVSVTAYASGPLASHLSPPNDIKSL	350
sp P26367 PAX6_HUMAN	-----GUYPG-----TSVPGP-----	172
	..***:*,*..	
gi 2133658 pir  I45557	ASIGHQRNCPVATEDIHLKELDGHQSDETGSGEGNSGGASNIQNTD	400
sp P26367 PAX6_HUMAN	-----TQDGCQQQEGG-----GENTINSISSNGEDSDE	199



