

# Data Integration using the Distributed Annotation System (DAS)

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Rob Finn, Stefan Gräf, David Jackson, Andreas Kähäri,  
Eugene Kulesha, Roger Pettett, James Smith,  
Jim Stalker, Tim J. P. Hubbard

- what is DAS
- what do we do with it
- DAS registration server
- latest developments



Ensembl v39 - Jun 2006

Search all Ensembl: Anything Go Help

**Use Ensembl to...**

- Run a BLAST search
- Search Ensembl
- Data mining [BioMart]
- Export data
- Download data

Docs and support

- Information
- What's New
- About Ensembl
- Ensembl data
- Software

**Other links**

- Home
- Sitemap
- Vega
- Pre Ensembl
- View previous release of page in Archive!
- Stable Archive! link for this page
- Archival sites
- Trace server

**What's New in Ensembl 39**

- Honeybee dropped from Ensembl (*Apis mellifera*)
- New Opossum assembly and genebuild (*Monodelphis domestica*)
- New Mouse assembly and genebuild (*Mus musculus*)
- New Ciona savignyi assembly and genebuild (*Ciona savignyi*)
- New Zebrafish assembly and genebuild (*Danio rerio*)

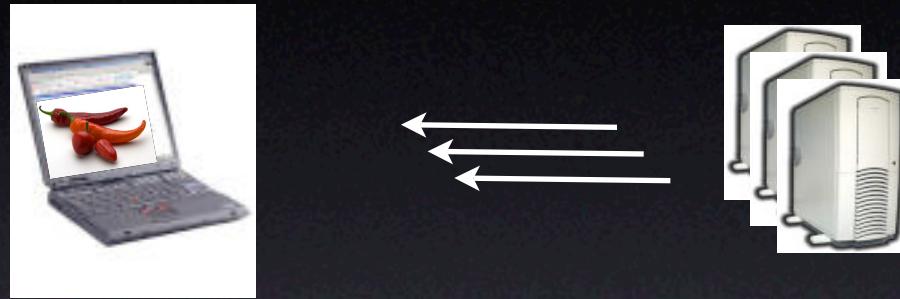
**Mammalian genomes**

- Homo sapiens* NCBI 36 | Vega
- Pan troglodytes* PanTro 1.0
- Macaca mulatta* Mmul 0.1 | pre!
- Mus musculus* UPDATED! NCBI m36 | Vega
- Rattus norvegicus* RNOH 3.4
- Oryctolagus cuniculus* RABBIT
- Canis familiaris* CanFam 1.0 | Vega | pre!
- Bos taurus* Btau 2.0
- Dasyurus novemcinctus* ARNIA
- Loxodonta africana* BROAD E1
- Echinops telfairi* TENREC

**Other species**

- Gallus gallus* VASHUC 1
- Xenopus tropicalis* JGI 4.1
- Danio rerio* UPDATED! zv 6 | Vega
- Takifugu rubripes* FGU34.0
- Tetraodon nigroviridis* TETRAODON 7
- Gasterosteus aculeatus* BROAD S1
- Ciona intestinalis* JGI 2
- Ciona savignyi* NEW! CSAV 2.0
- Drosophila melanogaster* BDGP 4
- Anopheles gambiae* AgamP3
- Aedes aegypti* AEDES 1

# Integration of personal data into bioinf. resources



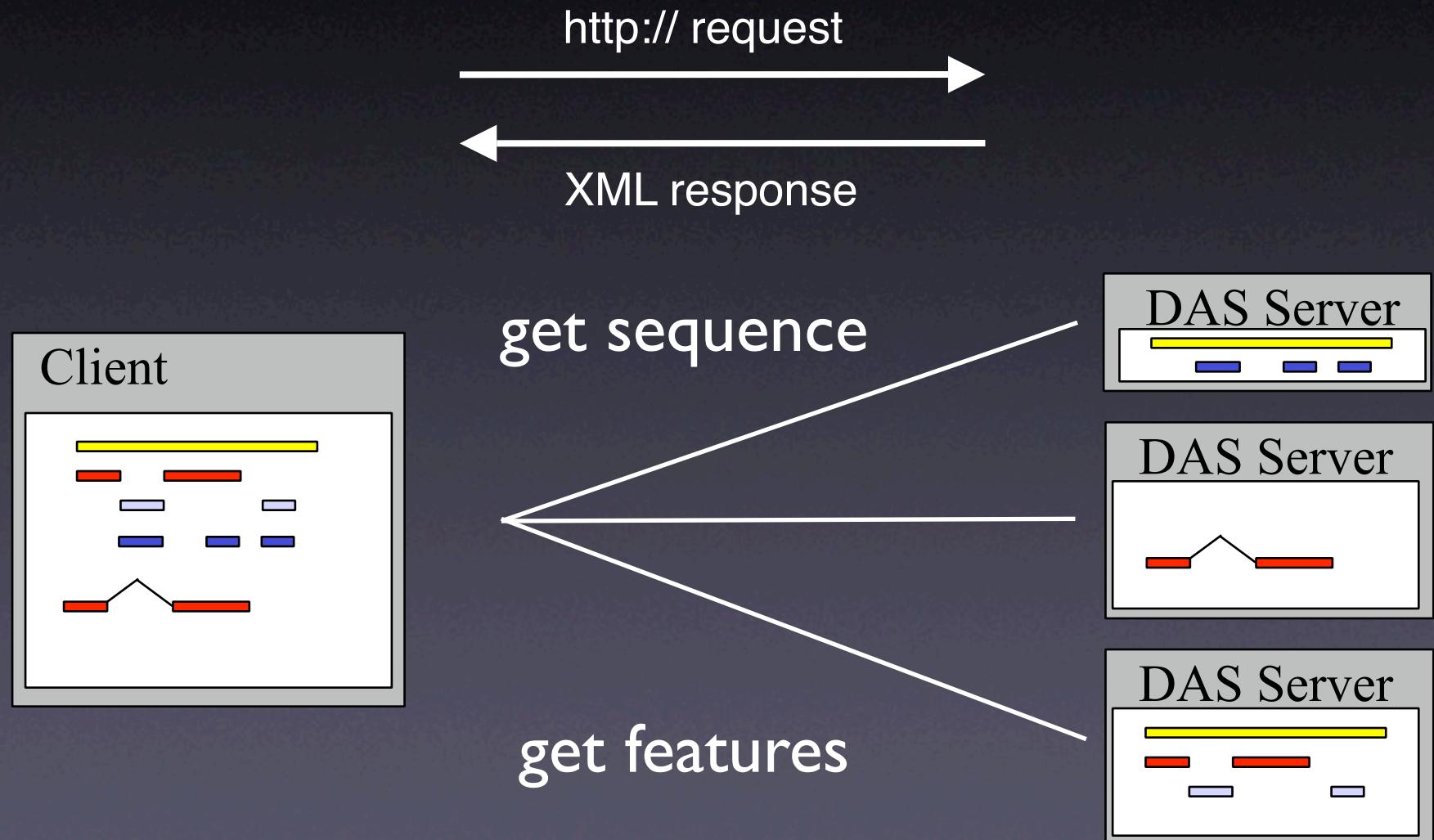
- Integration of annotations from external sources into local applications



- online access to most recent data versions
  - no need for local installations

# DAS, how it works

Dowell, Jokerst, Allen, Eddy, Stein  
BMC Bioinformatics 2001



# a few principles...

- Clients are “intelligent” (few)
- Servers are simple and easy to set up (many)
- (most of) data is precalculated
- libraries for server and client
- multiple programming languages

Ensembl v39 - Jun 2006

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## Other links

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- Stable Archive! link for this page
- Archive! sites
- Trace server



**Rattus norvegicus**  
3.4 update

## What's New in Ensembl 39

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- New Zebrafish assembly and genebuild (*Danio rerio*)

[More news...](#)

## About Ensembl

Ensembl is a joint project between EMBL-EBI and the Sanger Institute to develop a software system which produces and maintains automatic annotation on selected eukaryotic genomes. Ensembl is primarily funded by the Wellcome Trust.

This site provides [free access](#) to all the data and software from the Ensembl project. Click on a species name to browse the data.

Access to all the data produced by the project, and to the software used to analyse and present it, is provided free and without constraints. Some data and software may be subject to [third-party constraints](#).

For all enquiries, please [contact the Ensembl HelpDesk](#) ([helpdesk@ensembl.org](mailto:helpdesk@ensembl.org)).

## Other sites using the Ensembl system

EBI Genome Reviews database

## Mammalian genomes

***Homo sapiens***

NCBI 36 | Vega

***Pan troglodytes***

PanTro 1.0

***Macaca mulatta***MMUL 0.1 | *pre!****Mus musculus***

UPDATED! NCBI m36 | Vega

***Rattus norvegicus***RGSC 3.4 | *pre!****Oryctolagus cuniculus***

RABBIT

***Canis familiaris***CanFam 1.0 | Vega | *pre!****Bos taurus***

Btau 2.0

***Dasypus novemcinctus***

ARMA

***Loxodonta africana***

BROAD E1

***Echinops telfairi***

TENREC

***Monodelphis domestica***

UPDATED! MonDom 4

## Other species

***Gallus gallus***

WASHUC 1

***Xenopus tropicalis***

JGI 4.1

***Danio rerio***

UPDATED! 2y 6 | Vega

***Takifugu rubripes***

FUGU 4.0

***Tetraodon nigroviridis***

TETRAODON 7

***Gasterosteus aculeatus***

BROAD S1

***Oryzias latipes***

NEW! MEDAKA 1

***Ciona intestinalis***

JGI2

***Ciona savignyi***

UPDATED! CSAV 2.0

***Drosophila melanogaster***

BDGP 4

***Anopheles gambiae***

AgamP3

***Aedes aegypti***

AEDES 1

***Caenorhabditis elegans***

WS 150

***Saccharomyces cerevisiae***

SGD 1

- > 20 vertebrates / model organism
- 5 mill. page impressions / week
- 100 mirrors/internal installations worldwide
- open source
- used for other species as well
- MySQL
- 5-10 G / species + 100 G multi species data

Ensembl v39 - Jun 2006

Chromosome 12  
9,706,323 -  
0,559,649

View of Chromosome 12

Graphical view

Graphical overview

Export information about  
region

Export sequence as FASTA

Export EMBL file

Export Gene info in region

Export SNP info in region

Export Vega info in region

View data stored on  
another webserver in  
ContigView

View alignment with ...

View alongside ...

View Syntenic regions ...

View region in NCBI  
browser

View region in UCSC  
browser

Add Ensembl to...

Run a BLAST search

Search Ensembl

Data mining [BioMart]

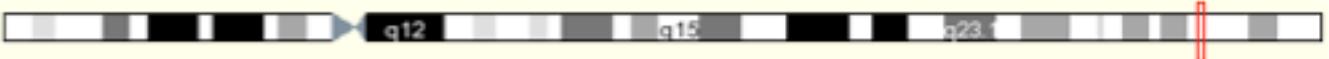
Upload and view data

chromosome

Export data

## Chromosome 12

Chr. 12



## Overview

## Detailed view

Features ▾ Comparative ▾ DAS Sources ▾ Repeats ▾ Decorations ▾ Export ▾ Image size ▾ Help ▾

Jump to region  :  CPG island clones  
 Compugen oligos  
 DECIPHER  
 Human SPG predictions  
 Human geneid predictions  
 IMGT Genes  
 Illumina probes  
 NCBI Gnomon  
 OMIM Gene map  
 OMIM Morbid map  
 PDB\_Splice  
 RZPD expression clones  
 RZPD gene silencing  
 RZPD verified cDNA  
 RefSeq  
 Sanger Hver Array  
 Toronto Poly  
 Toronto Poly [loc]  
 RZPD verif. cDNA  
 Manage sources...  
 URL based data...  
 EMBL mRNAs

Refresh Band:  Refresh

Zoom Window > 1MB > 2MB > 5MB >>

Chr. 12 Length For

120.00 Mb 853.33 Kb 120.20 Mb 120.40 Mb

R:ENSG00000135100 RZPD:cDNA:ENSG00000135124  
RZPD:cDNA:ENSG0000089041 RZPD:cDNA:ENSG00000170633 RZPD:cDNA:ENSG00000170633  
RZPD:PEC:ENSG0000089041 RZPD:PEC:ENSG00000170633  
RZPD:PEC:ENSG00000135124 RZPD:PEC:ENSG00000135124

ENSG00000135100

GENSCAN0000017712 > Ah-initio Genscan trans

CLOSE MENU

Add your own  
uses Registry

- [Transcript information](#)
- [Exon information](#)
- [Protein information](#)
- [Export protein data](#)

Chromosome 17  
74,183,768 - 74,289,900

- [View of Chromosome 17](#)
- [Graphical view](#)
- [Graphical overview](#)
- [Export information about region](#)
- [Export sequence as FASTA](#)
- [Export EMBL file](#)
- [Export Gene info in region](#)
- [Export SNP info in region](#)
- [Export Vega info in region](#)

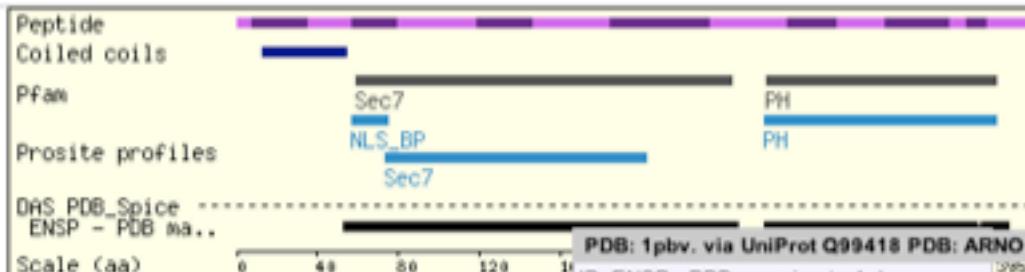
#### Use Ensembl to...

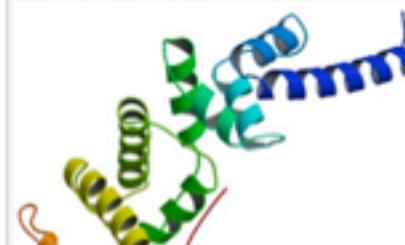
- [Run a BLAST search](#)
- [Search Ensembl](#)
- [Data mining \[BioMart\]](#)
- [Upload your own data](#)
- [Export data](#)
- [Download data](#)

#### Docs and downloads

- [Information](#)
- [What's New](#)
- [About Ensembl](#)
- [Ensembl data](#)
- [Software](#)

# Linking protein structure to e! Peptide view

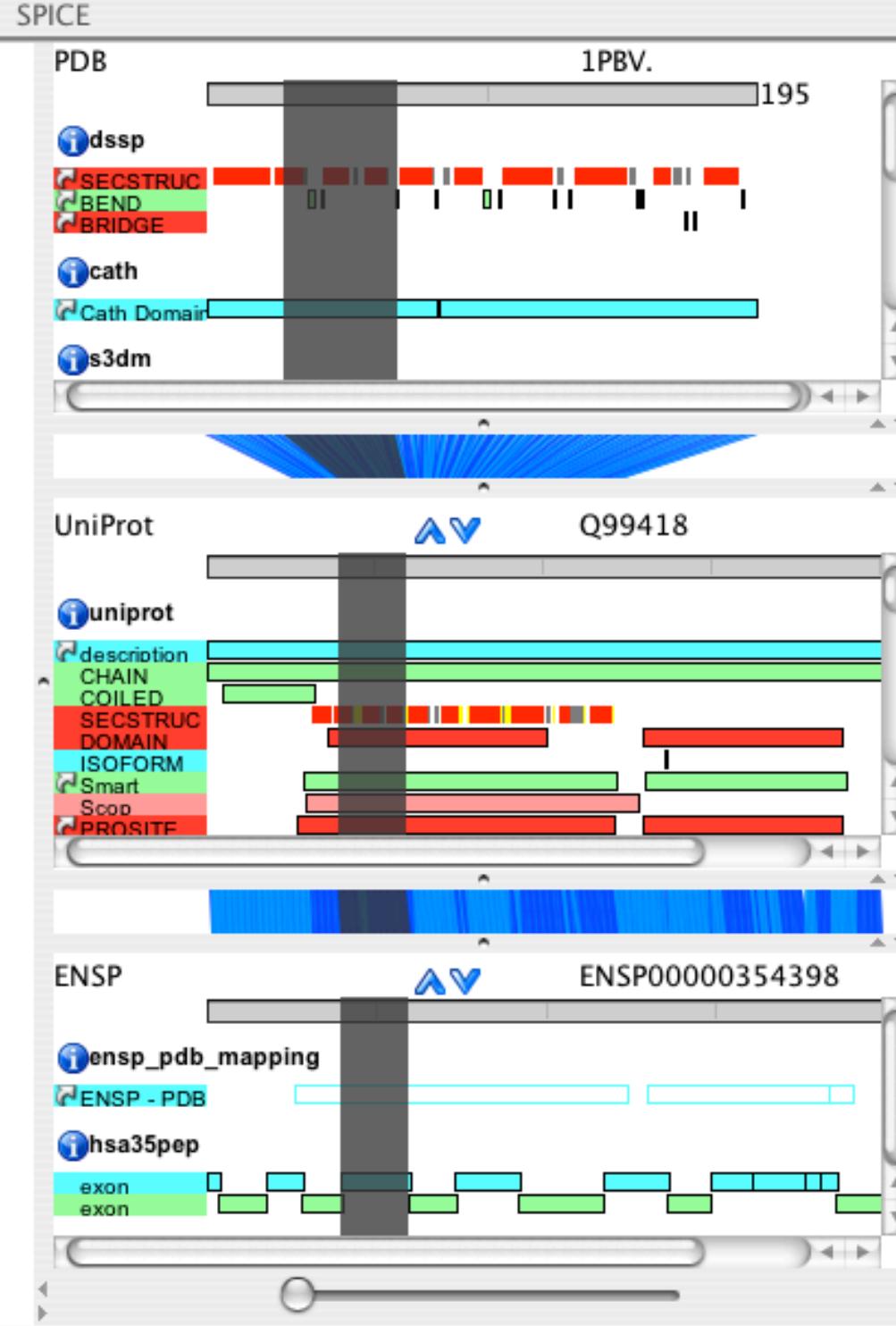
Prediction Method	Genes were annotated by the Ensembl automatic analysis pipeline using either a GeneWise/Exonerate model from a database protein or a set of aligned cDNAs followed by an ORF prediction. GeneWise/Exonerate models are further combined with available aligned cDNAs to annotate UTRs (For more information see V.Curwen et al., Genome Res. 2004 14:942-50.)
InterPro	<a href="#">IPR001472</a> Bipartite nuclear localization signal - [ <a href="#">View other genes with this domain</a> ] <a href="#">IPR000904</a> SEC7-like - [ <a href="#">View other genes with this domain</a> ] <a href="#">IPR001849</a> Pleckstrin-like - [ <a href="#">View other genes with this domain</a> ]
Protein Family	<a href="#">ENSF00000001251</a> : CYTOHESIN This cluster contains 4 Ensembl gene member(s)
Protein Features	 <div style="border: 1px solid black; padding: 5px; margin-top: 10px;">       PDB: 1pbv. via UniProt Q99418 PDB: ARNO               ID: ENSP - PDB mapping to 1pbv.               TYPE: ENSP - PDB mapping               METHOD: Compara, MSD-Mapping of UniProt to PDB               LINK: <a href="http://das.sanger.ac.uk/registry/showspice.jsp?pdb=1pbv">http://das.sanger.ac.uk/registry/showspice.jsp?pdb=1pbv</a>         NOTE: PDB: 1pbv. 85% id 49% coverage via UniProt Q99418 PDB:ARNO     </div>
Protein Sequence	<pre>MEEDISTYPSDLNREIPIQILEHNRKQKILLADIQRIKDIIAEVVNEIIHLGSTEETKHNH QPKQKQHGRKKHNPDKKGIGELIINLLIANTCEDIQFLIDKSEGILHKWIGDYLGEK&gt; EINIOQHIGLIVELMEITISHLVWQLPQTLASTRLPGEIAKQIDRMTHETAQRTCOCHNGVY QSTDTCVYLSTAIIMHTS1JNPNVKKPTVERIIPHNGINDGGDLPITLLPHLYESIK HEPEKIPEDDGDHLTWTTHPDREGWLIKLGGRVKTWNQMTILTWCYLTYTEYTTKIKP RGIIPLHLHSIREVEIKKKPQHCFELYIPSHKDQVQKACKTERDGKVVIGHHFTGSQLQK&gt; EKEEWIKCKRAISRKTTIMEAKKKVSYTKPH</pre>
	Exon alternating text colour      Residue overlap splice site



# SPICE browser

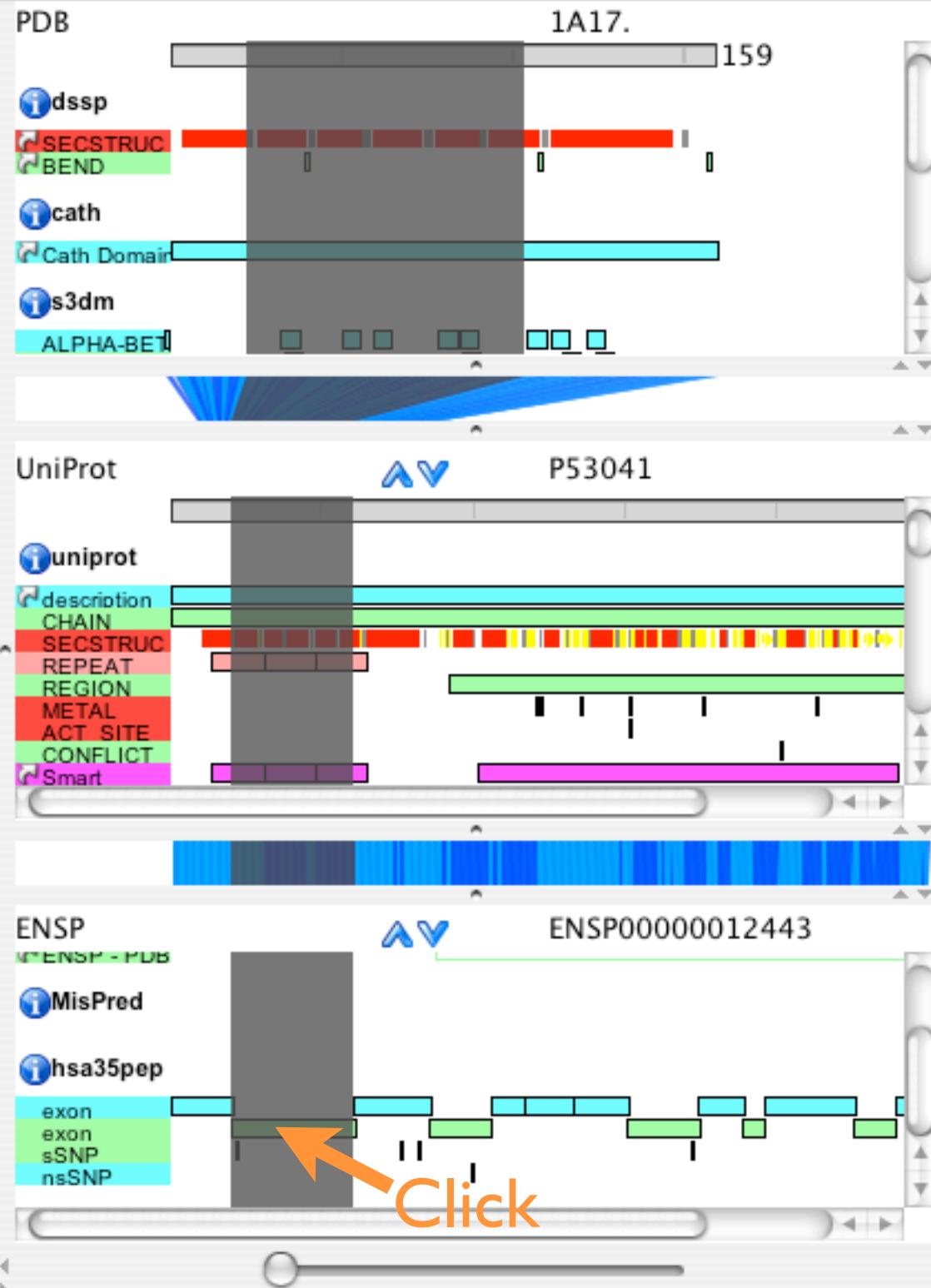


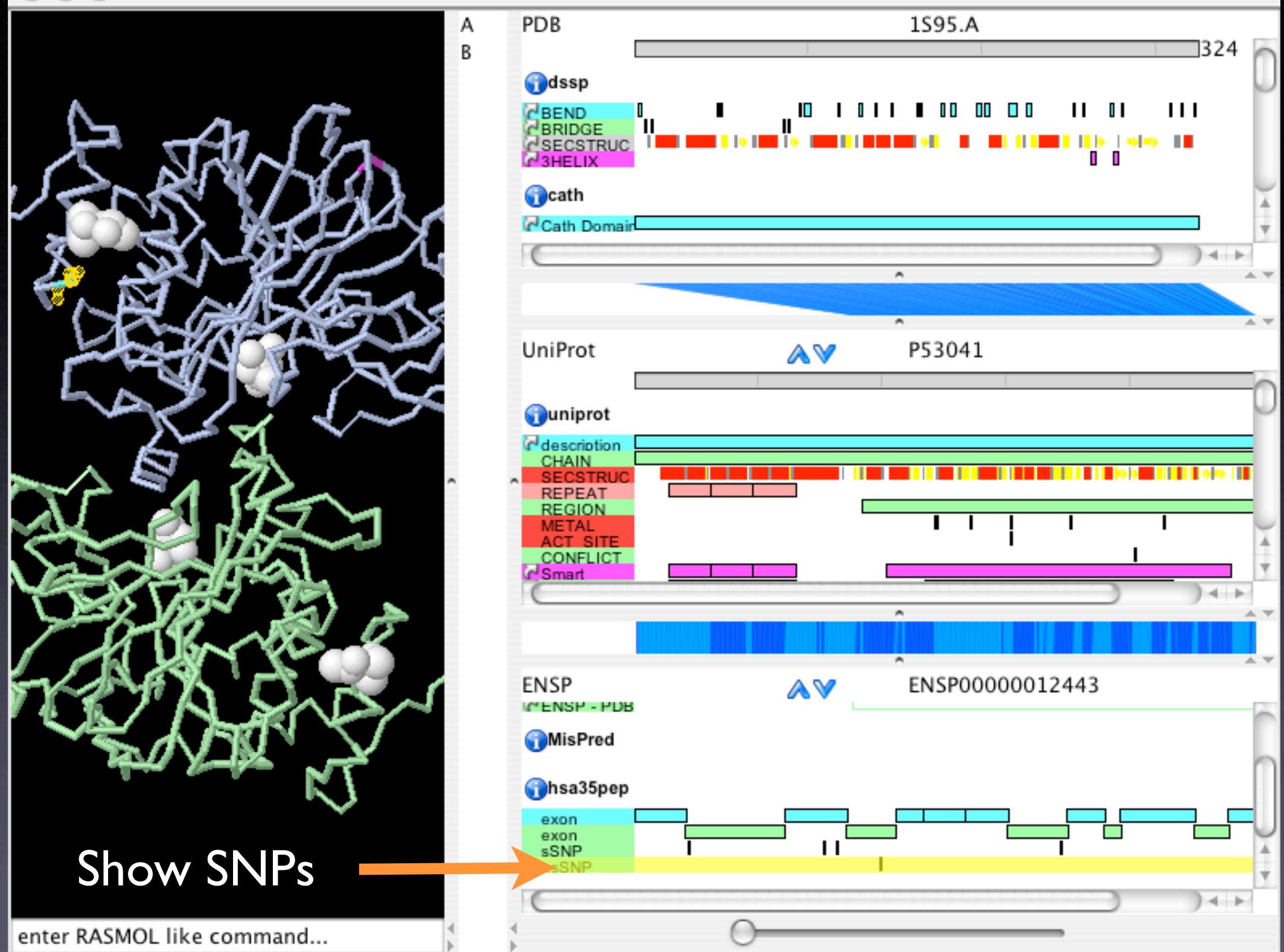
enter RASMOL like command...



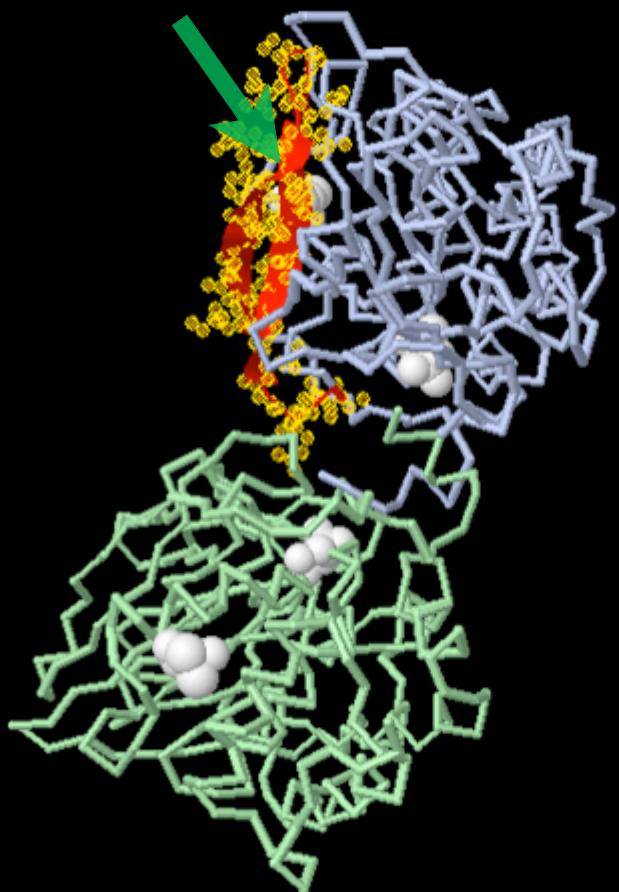


enter RASMOL like command...





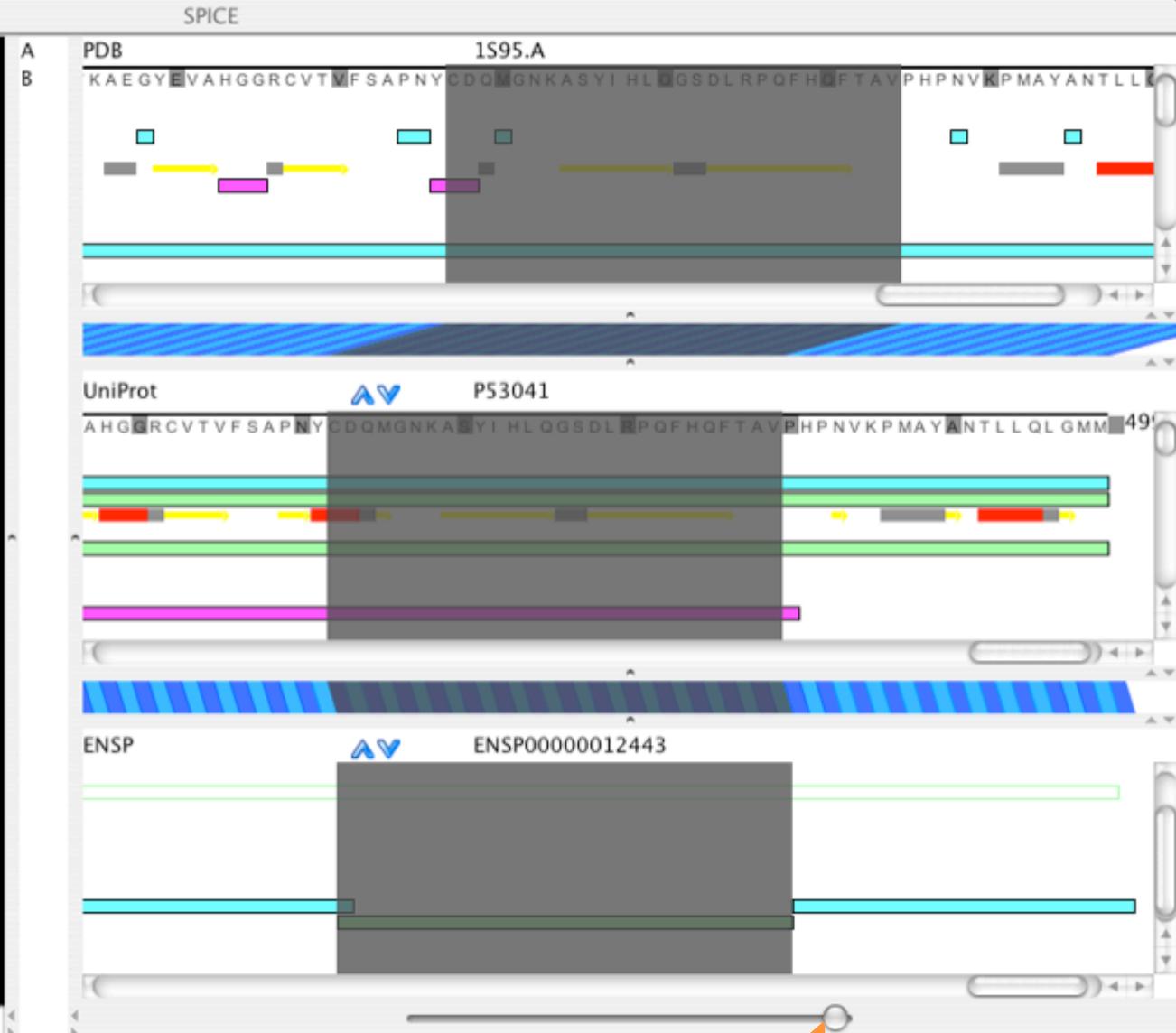
# interact with Menu & RASMOL



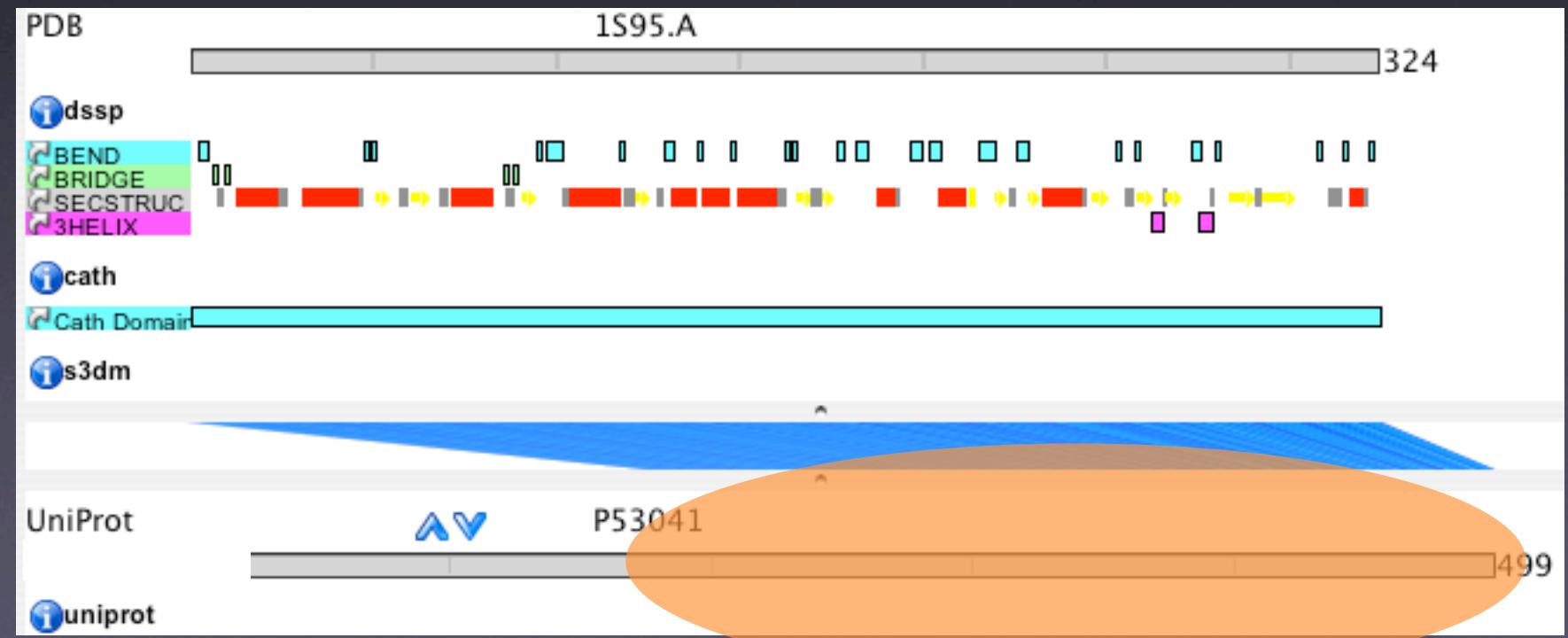
enter RASMOL like command:

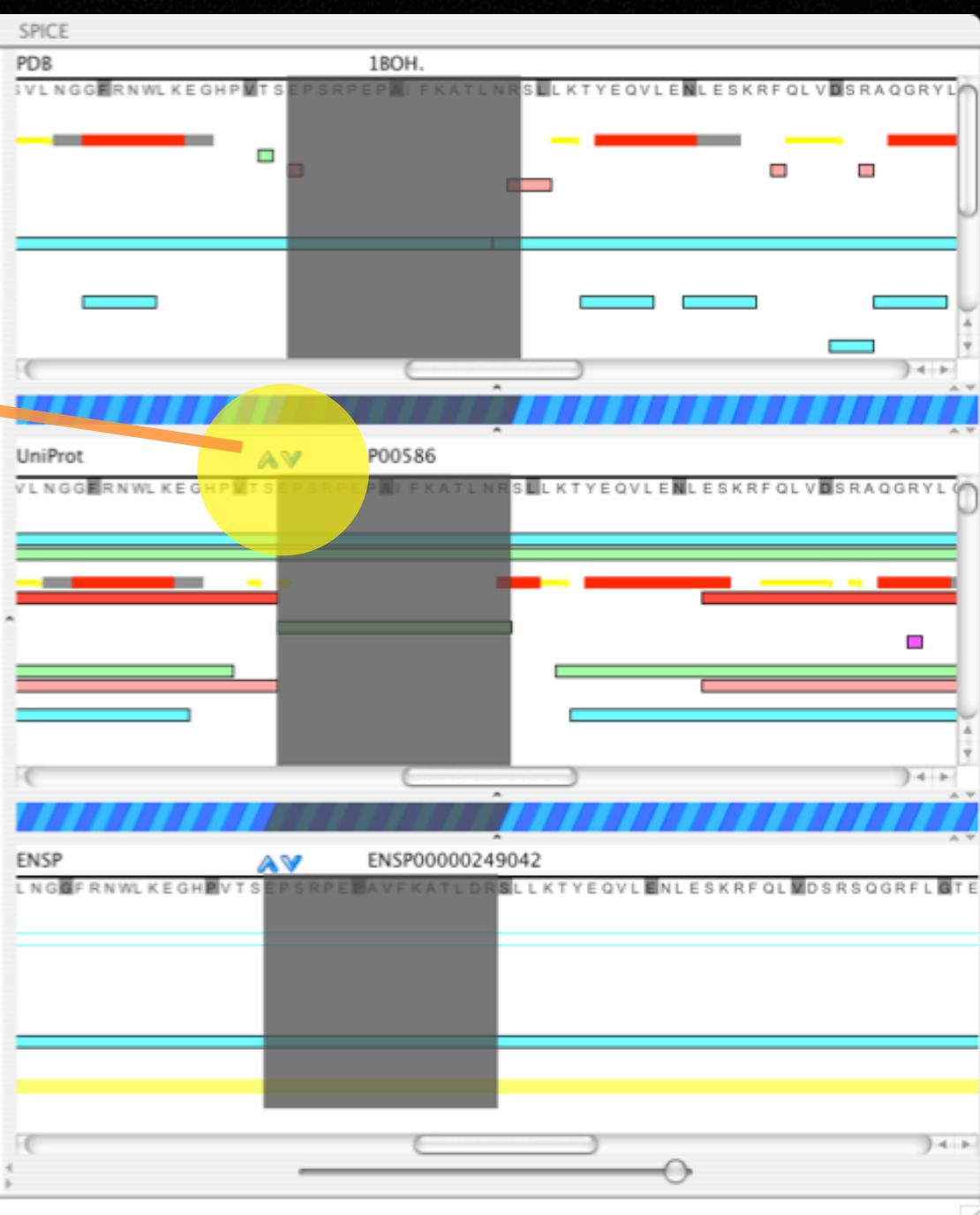
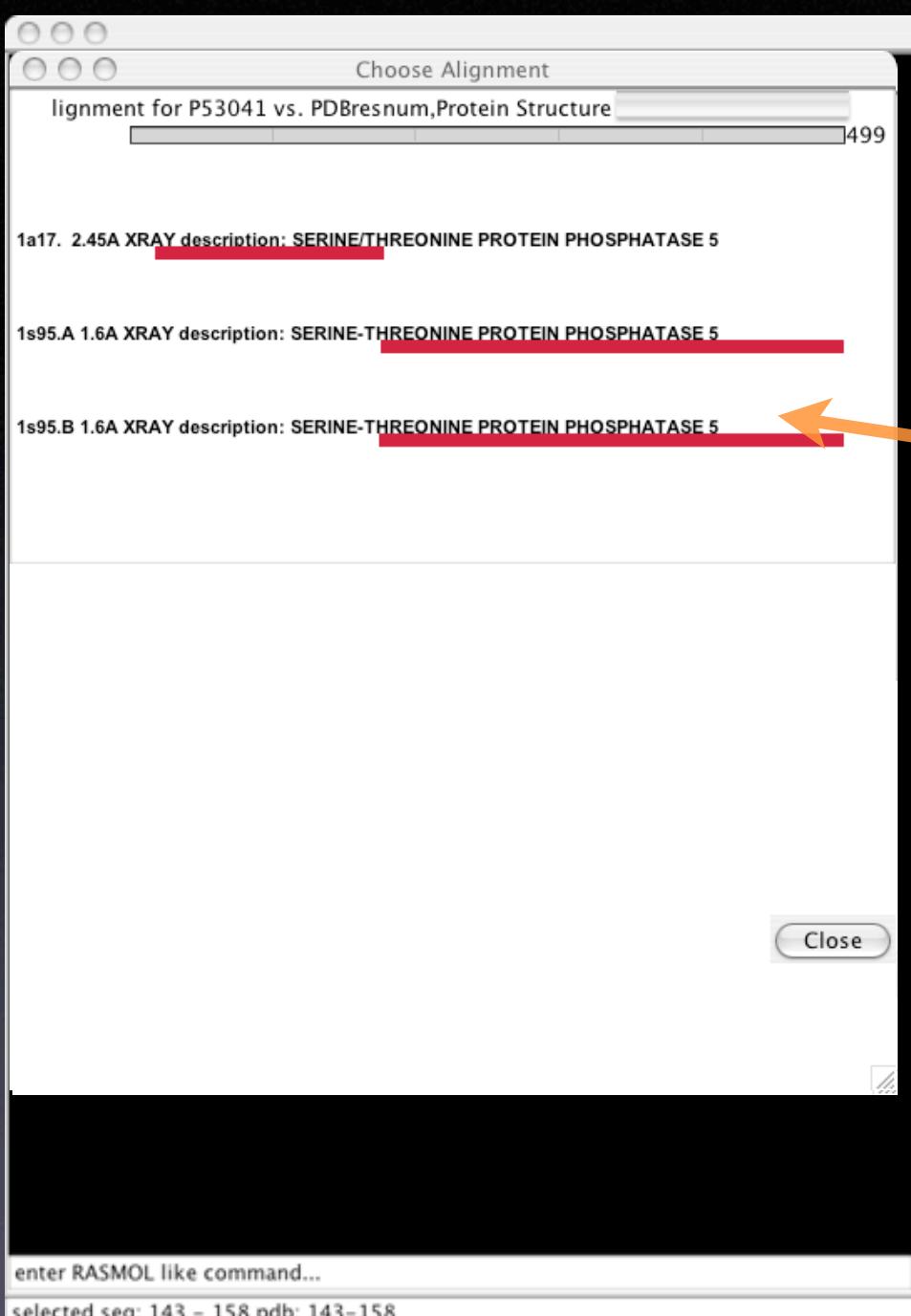
selected seq: 277 - 304 pdb: 479

RASMOL  
commands



Zoom





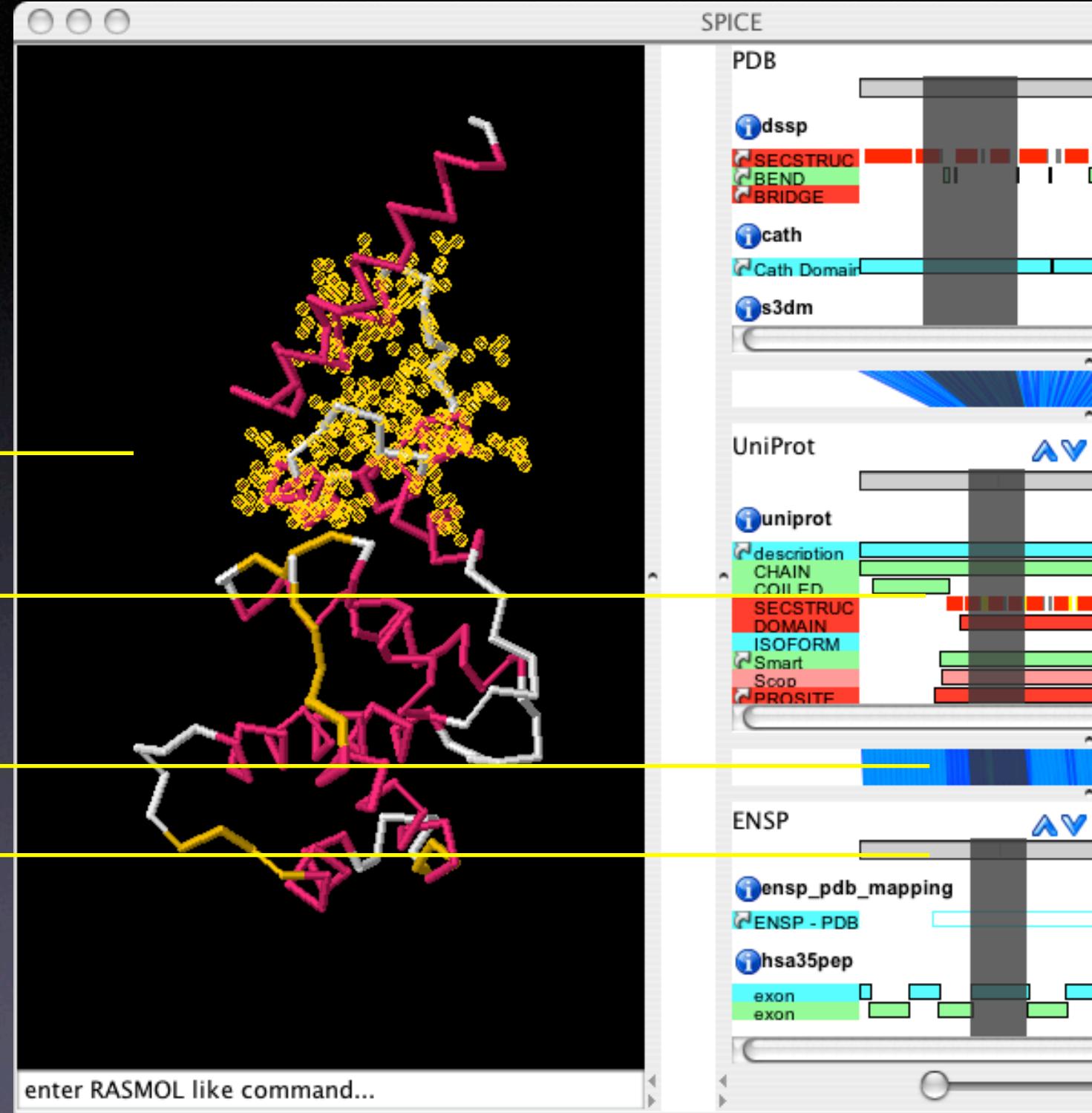
# DAS commands

Structure

Features

Alignment

Sequence



The screenshot shows the Ensembl homepage from June 2006. At the top, there's a search bar with the placeholder "Search all Ensembl: Anything". Below the search bar, there are several sections:

- Use Ensembl to...** includes links for running a BLAST search, searching Ensembl, data mining (BioMart), exporting data, and downloading data.
- Docs and downloads** includes links for information, what's new, about Ensembl, ensemble data, software, and other links like Home, Sitemap, Vega, and Trace server.
- What's New in Ensembl 39** lists recent updates: Honeybee dropped from Ensembl (*Apis mellifera*), New Opossum assembly and genebuild (*Monodelphis domestica*), New Mouse assembly and genebuild (*Mus musculus*), New Ciona savignyi assembly and genebuild (*Ciona savignyi*), and New Zebrafish assembly and genebuild (*Danio rerio*).
- About Ensembl** provides a brief overview of the project, mentioning it is a joint project between EMBL-EBI and the Sanger Institute, and links to the Welcome Trust.
- Mammalian genomes** lists species: *Homo sapiens* (NCBI 36 | Vega), *Pan troglodytes* (PanTro 1.0), *Macaca mulatta* (Mmul 0.1 | pre!), *Mus musculus* (UPDATED! NCBI m36 | Vega), *Rattus norvegicus* (Rnor 5.4), *Oryctolagus cuniculus* (RABBIT), *Canis familiaris* (CanFam 1.0 | Vega | pre!), *Bos taurus* (Btau 2.0), *Dasyurus novemcinctus* (AR91A), *Loxodonta africana* (BROAO-E1), and *Echinops telfairi* (TENREC).
- Other species** lists species: *Gallus gallus* (VASHUC 1), *Xenopus tropicalis* (J014.1), *Danio rerio* (UPDATED! zv 6 | Vega), *Takifugu rubripes* (FUSU 4.0), *Tetraodon nigroviridis* (TETRAODON 7), *Gasterosteus aculeatus* (BROAD S1), *Ciona intestinalis* (J012), *Ciona savignyi* (NEW! CSAV 2.0), *Drosophila melanogaster* (EDGP 4), *Anopheles gambiae* (AgamP3), and *Aedes aegypti* (ADES 1).

## Java Web Start

auto install  
latest version  
send arguments

DAS registry

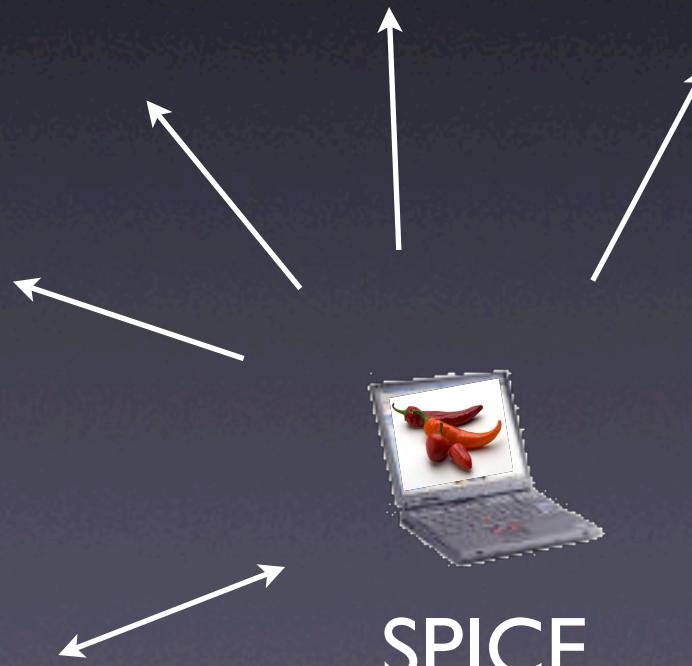


Meta information  
about DAS servers

DAS registry



SPICE



# The DAS registration server

DS_110	  	dssp	cmbi4.cmbi.ru.nl/das/dssp/	<a href="#">features</a>	PDBresnum,Protein Structure
DS_111	  	cath	cathwww.biochem ... 0/das/cath_pdb/	<a href="#">features</a>	PDBresnum,Protein Structure
DS_112	  	structure	das.sanger.ac.uk/das/structure/	<a href="#">structure</a>	PDBresnum,Protein Structure
DS_113	  	alig_pdb_sp	das.sanger.ac.uk/das/msdpdbsp/	<a href="#">alignment</a>	UniProt,Protein Sequence PDBresnum,Protein Structure
DS_114	    	signalp	genome.cbs.dtu.dk:9000/das signalp/	<a href="#">types</a> <a href="#">features</a>	UniProt,Protein Sequence
DS_115	    	netphos	genome.cbs.dtu.dk:9000/das/netphos/	<a href="#">types</a> <a href="#">features</a>	UniProt,Protein Sequence
DS_116	    	netoglyc	genome.cbs.dtu.dk:9000/das/netoglyc/	<a href="#">types</a> <a href="#">features</a>	UniProt,Protein Sequence
DS_117	    	tmhmm	genome.cbs.dtu.dk:9000		

<http://das.sanger.ac.uk/registry/>

# DAS registration server

- allows to “publish” DAS servers & share with community
- communicates with clients
- regularly checks servers, sends notification

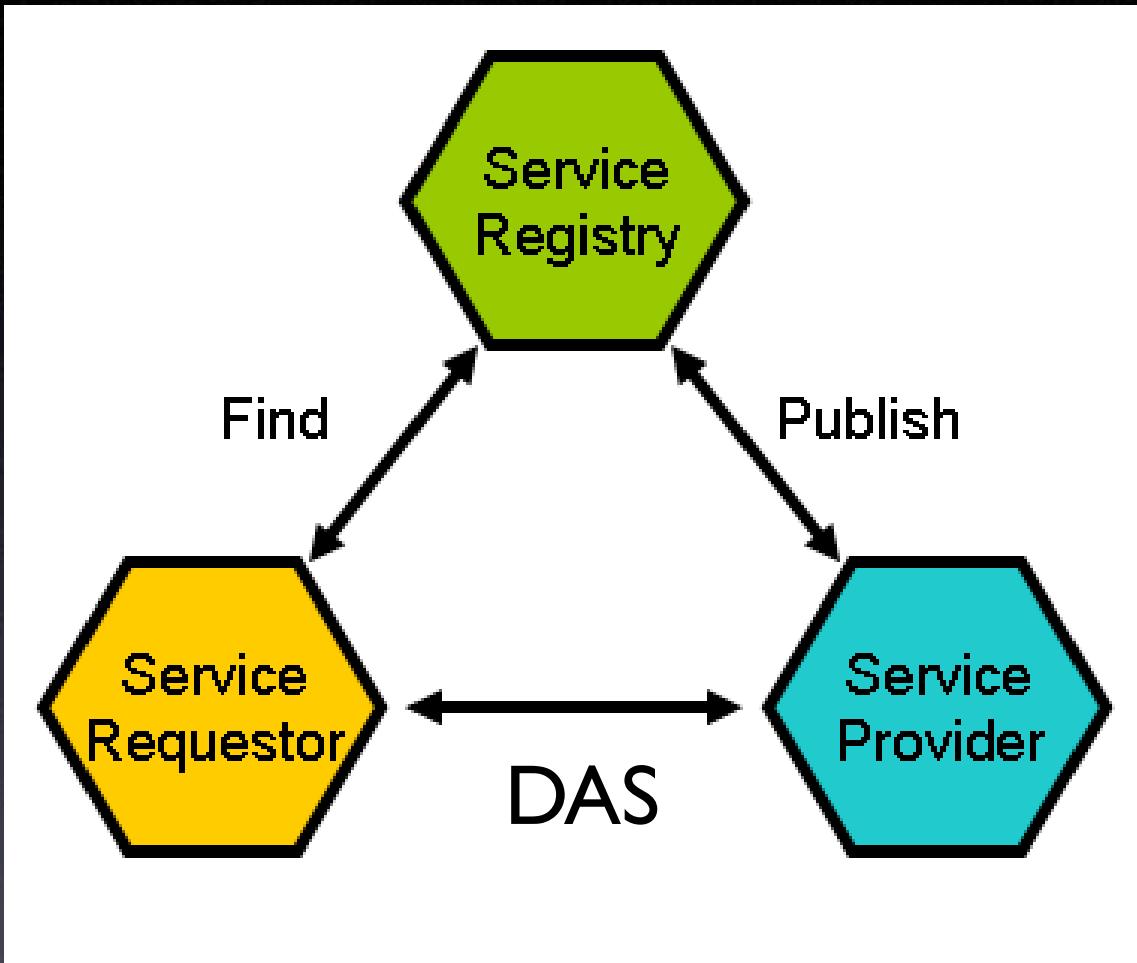
# What is the glue?

- “Coordinate Systems”
  - Authority
  - Type of data
  - Version
  - Organism (optional)

# Clients and Coordinate Systems

- Ensembl - most of the views can display DAS sources from multiple CS
- SPICE - PDB, UniProt, Ensp
- Dasty - UniProt

# DAS registration server

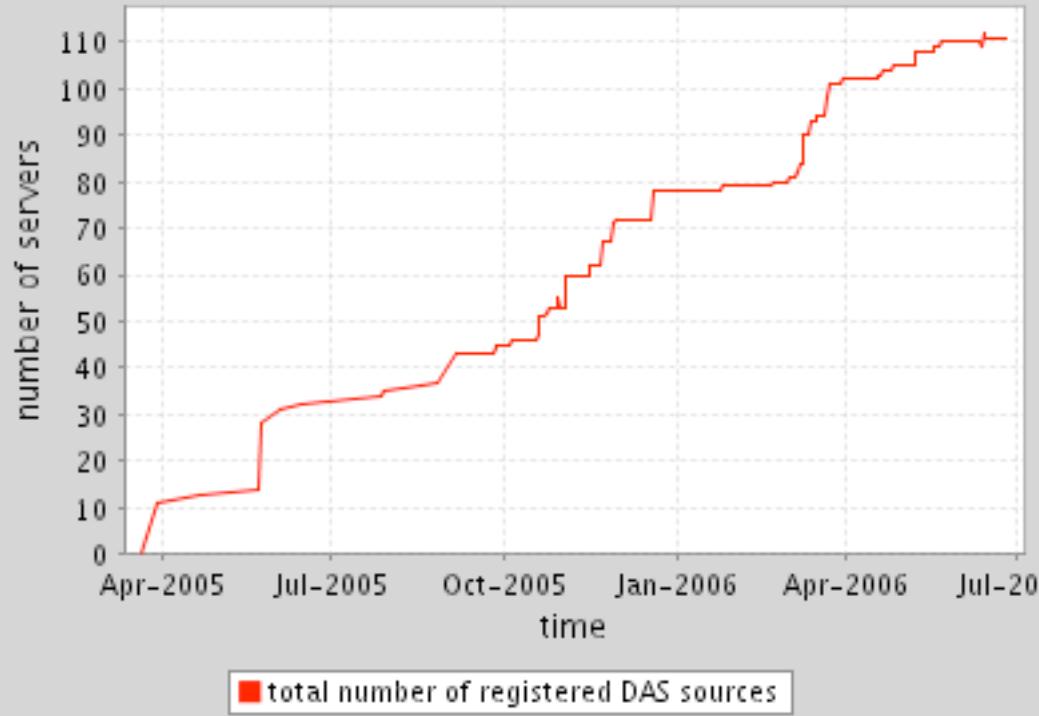


e.g.  
Ensembl,  
SPICE

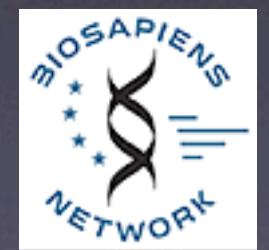
a DAS source

the DAS - SOA

## registered DAS servers over time



111 DAS sources  
26 institutions  
12 countries



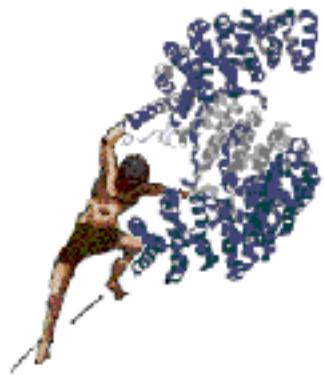
+ others

# DAS - issues

- inconsistent implementations
- no consistent use annotation types
- error handling
- searches not possible - in DAS/I
- open sharing of data - low security

# SISYPHUS

Submit Query



## Structural alignments for proteins with non-trivial relationships

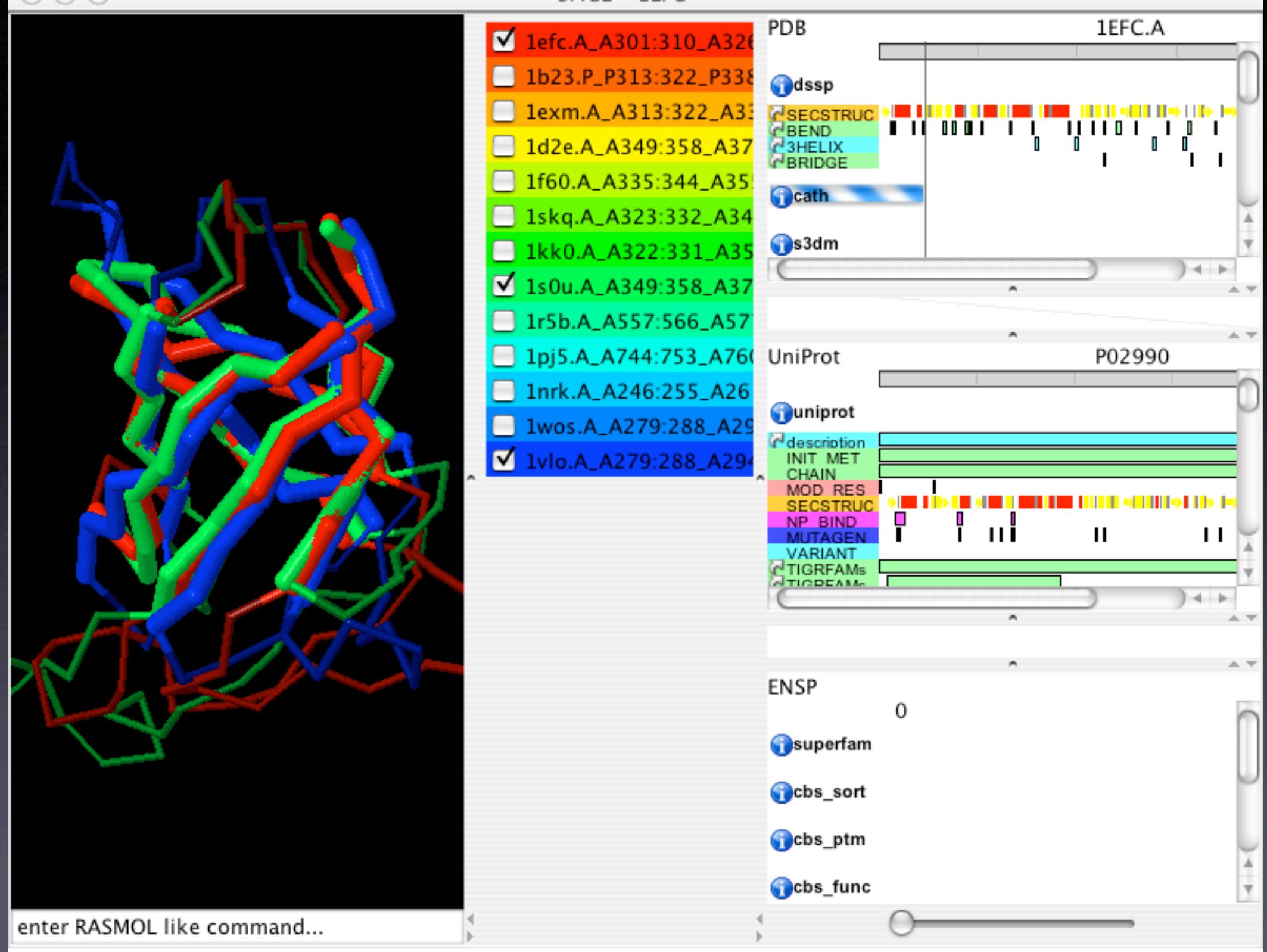
*Sisyphus: in greek: crafty*

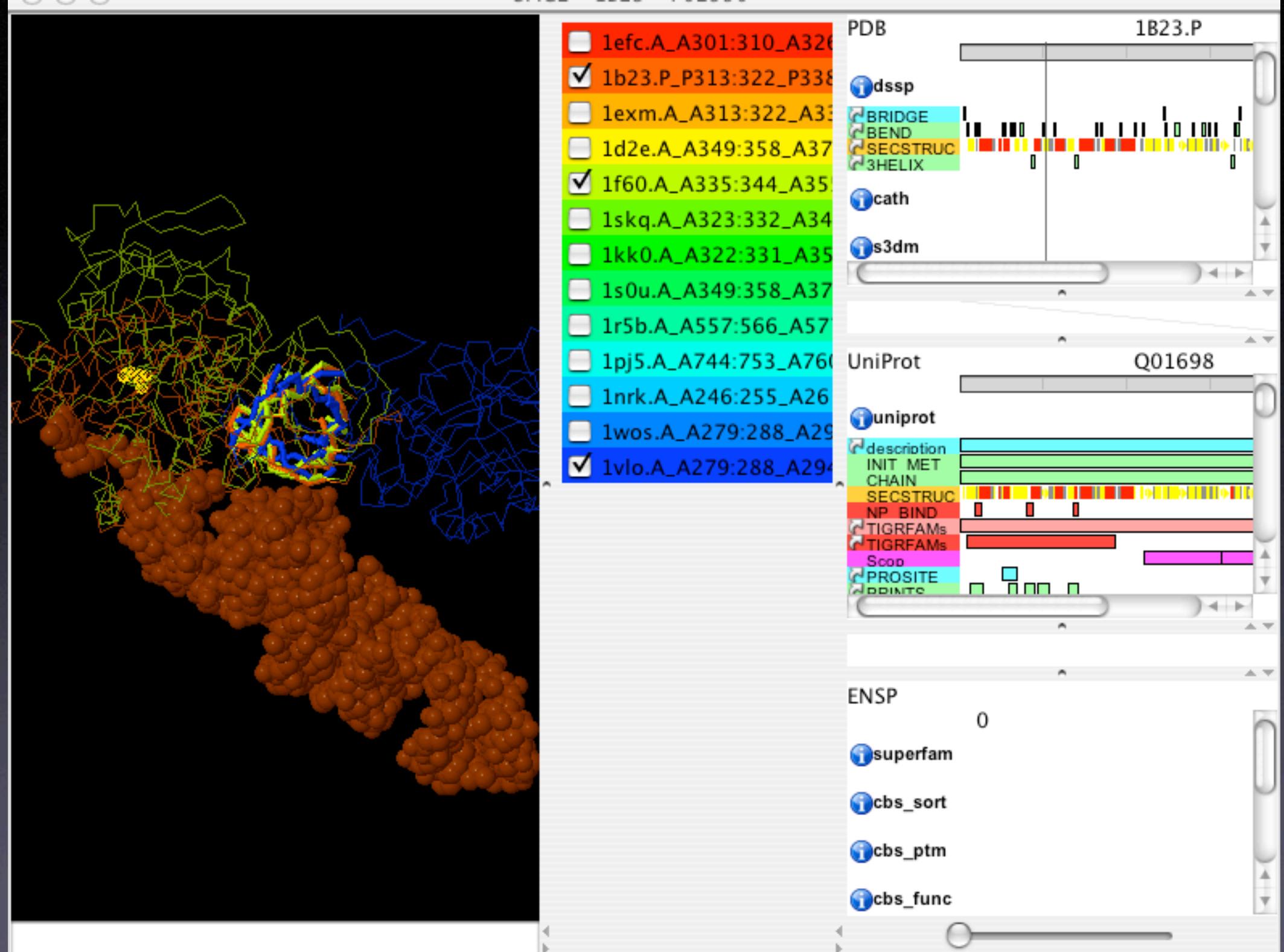
### Authors:

Antonina Andreeva, Andreas Prlic, Tim Hubbard, Alexey Murzin

```
SATVGIFVDAgsraenvknNGTAHFLEHLAFkgtqnr-----pqQGIELeienig---shLNAYTsrs----eNTVYYAKSLq--edIPKAVDIV  
TTHIAIALEgvswsap--dYFVALATQAIVGnwdraigtgtn-----spSPLAVaaasqngslansYMSFStsyadsgLWGMYIVTDsnehnVRLIVNE  
FSALGLYIDagsrfegrnlKGCTHILDRLAFkstehv-----egRAMAEtellelg---gnYQCTSsr----eNLMYQASVFn--qdVGKMLQL  
LFHIQIGFEglpidhp--dIYALATLQTLLGgggsfsaggpgkgmystslyTHVLNqy----yfvenCVAFNhsysdsgIFGISLSCIp--qaAPQAVEV  
TCTVGWVIDagsryeseknNGAGYFVEHLAFkgtknr-----pgNALEKevesmg---ahLNAYStr----eHTAYYIKALS--kdLPKA  
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LAHVAIAVEgpgwahp--dLVALQVANAIIGHydrtyggglh-----ssSPLASiavtn-klcqsFQTFSicysetgLFGFYFVCDrms--IDDMMFV  
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ISTLAVKVHggsryat--kDGVAHLLNRNFqntntr-----saLKLVResselg---gtFKSTLdr----eYITLKATFLk--ddLPYYVNA  
DSVAAIGIPvn---ka--sLAQYEVLANLYLsal-----SELSGli-----SSAKLdkftdgglFTLFVRDQ-dsavVSSNIKK  
LVHAAIVAEsaaiigga--eANAFSVLQHVLG-----ANPHVkrnp----fdVSAFNasysdsgLFGFYTISQaay--AGQVIKA  
GSTIGVFIKagsryenssnLGTSHLLRLASSlttkga-----ssFKITRgieavg---gkLSVEStr----eNMAYTVECLr--ddVEILMEF  
ASRIGLFIKagsryensnnLGTSHLLRLASSlttkga-----ssFKITRgieavg---gkLWYr----eNMAYTVECLr--ddVDILMEF  
LVHAALVAEsaaiagsa--eANAFSVLQHVLGagphvkrgsna-----tSSLYQavakgvhapioQasvsdsSEVTISQa--eSDVIIKA
```

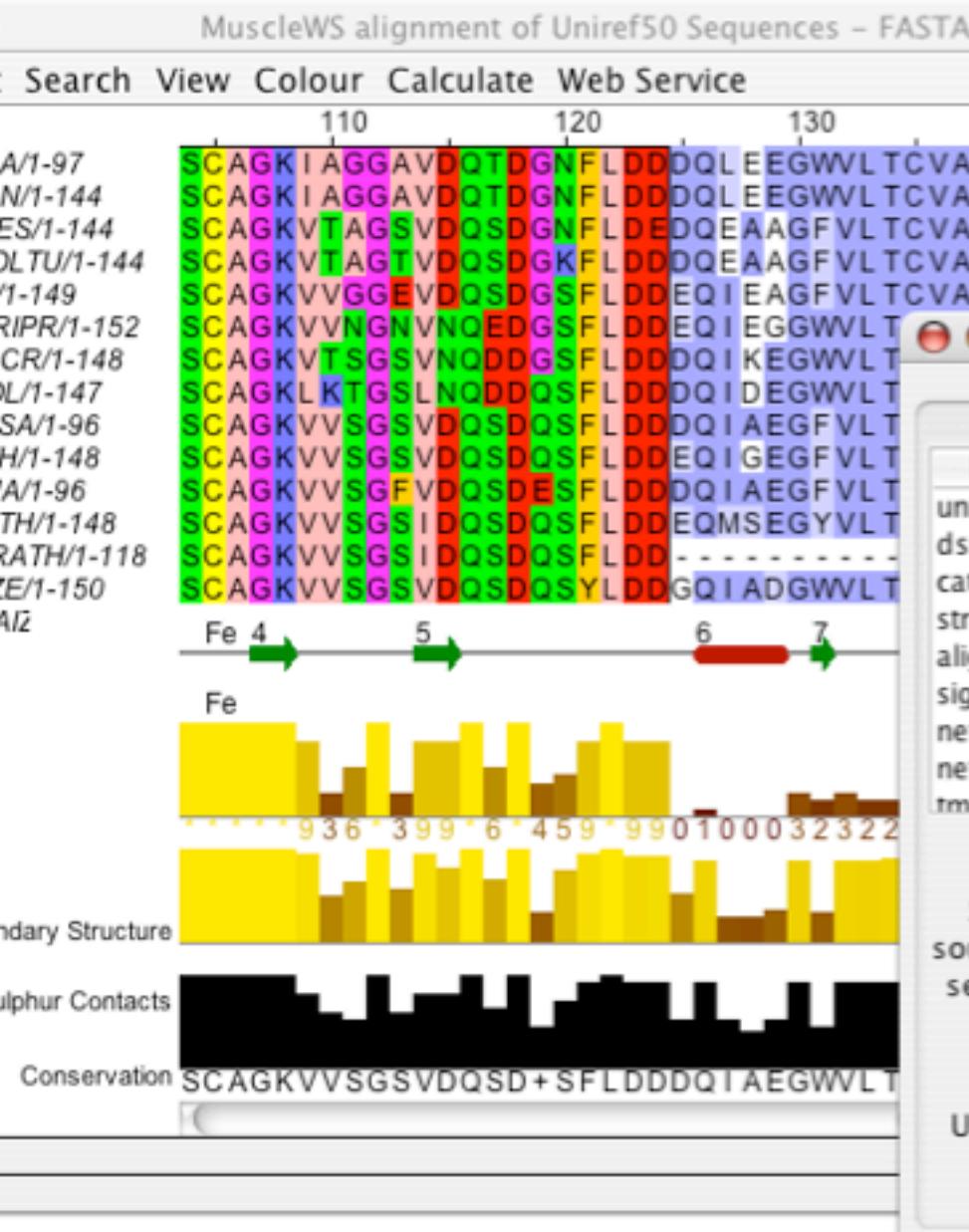






- Alignment DAS:
- rotation matrices, shift vectors
- range information (optional)

## Help Window



Average distance tree ...

File View

Preferences

DAS Settings

Nickname	Use Source
uniprot	<input checked="" type="checkbox"/>
dssp	<input type="checkbox"/>
cath	<input type="checkbox"/>
structure	<input type="checkbox"/>
alig_pdb_sp	<input checked="" type="checkbox"/>
signalp	<input checked="" type="checkbox"/>
netphos	<input checked="" type="checkbox"/>
netoglyc	<input type="checkbox"/>
tmhmm	<input checked="" type="checkbox"/>

Select a DAS service from the table to read a full description here

Show sources serving using coordinates

All Protein Sequence Protein Structure Gene\_ID Chromosome

All UniProt PDBresnum Ensembl ZV4 MGI

Use Registry <http://www.spice-3d.org/dasregistry/das1/sources>

Refresh Available Sources Add Local Source

# Acknowledgments

- T. Down, T. Hubbard
- Web Team, E. Kulesha, R. Pettett, T. Cox
- eFamily Project
- S. Gräf, A. Kahari, BioSapiens
- A. Murzin, A. Andreeva
- R. Finn, H. Hotz, A. Ahmed
- Jmol, Biojava, MSD, everybody who sets up DAS servers