

- 9.00-9.15am.** **Overview of the day**
- 9.15am - 10.30am.** **Session 1. Introduction to Jalview**
- starting the application, importing alignments, basic editing and creating figures.
- 10.30-11am.** **Coffee**
- 11am – 11.30am** **Geoff Barton: Multiple alignment and Analysis**
- 11.30am - 12.30pm.** **Session 2: Alignment & alignment analysis**
- Creating sequence alignments, importing and calculating trees, tree based alignment analysis
- 12.30pm to 1.30pm.** **Lunch**
- 1.30pm – 3.00pm.** **Session 3: Annotating sequences & alignments**
- Creating and viewing sequence annotation
 - Protein Secondary structure prediction
- 3.00pm – 3.30pm** **Coffee**
- 3.30pm – 4.30pm.** **Session 4: Working with molecular structures**
- Viewing 3D Structures, superimpositions, mapping disorder and alignment quality
 - Viewing RNA Secondary Structure
- 4.30pm – 4.45pm.** **Wrapup – what we didn't cover today**

Course materials

Everything is online

<http://www.jalview.org/tutorial/training-materials/2014/London/Dec/>

- These slides**
- Jalview v2.8.2 Manual (v1.5.0)**
 - Log in and**
 - Open the manual in your PDF Viewer NOW**

- Session 4
 - Working with structures
 - Viewing PDB data
 - Superimposing structures
 - Mapping data onto structure
 - Disorder prediction
 - RNA helix shading

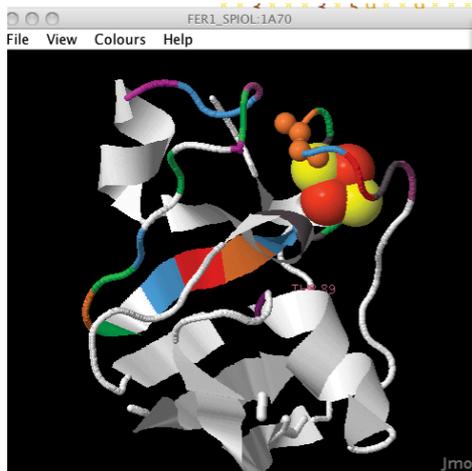
Desktop Structure Visualization

3D structures and 2D RNA diagrams



Jmol

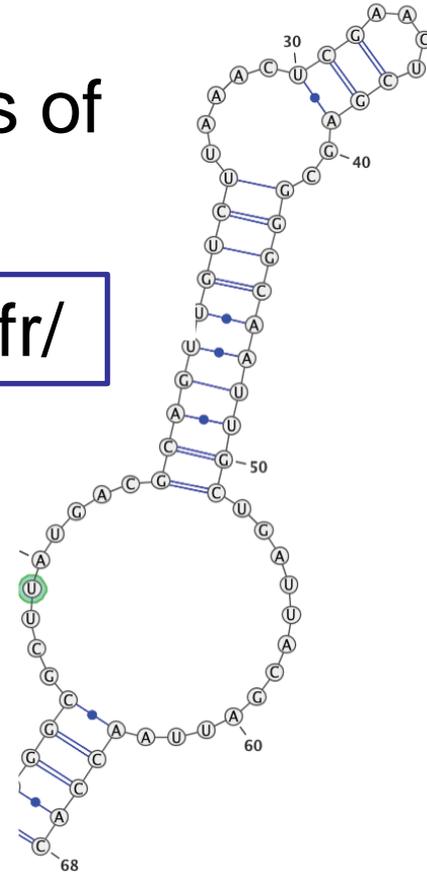
<http://jmol.sourceforge.net/>

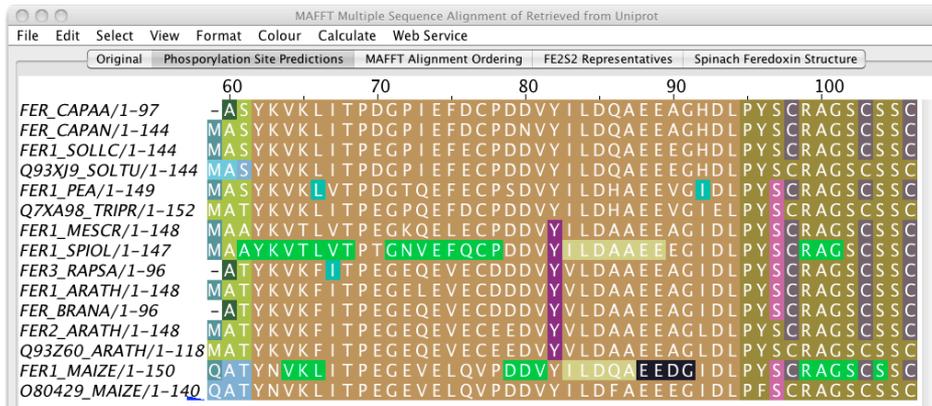


VARNA

Visual Analysis of
RNA

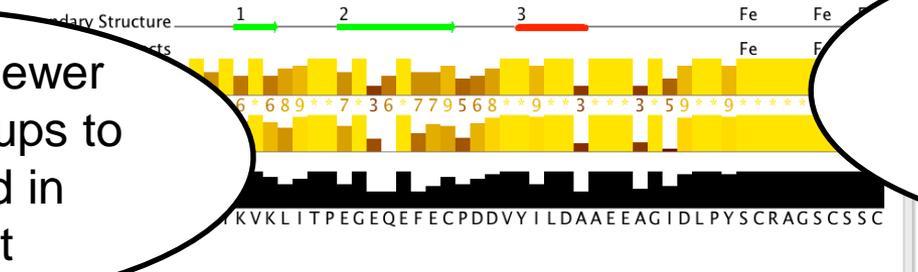
<http://varna.lri.fr/>



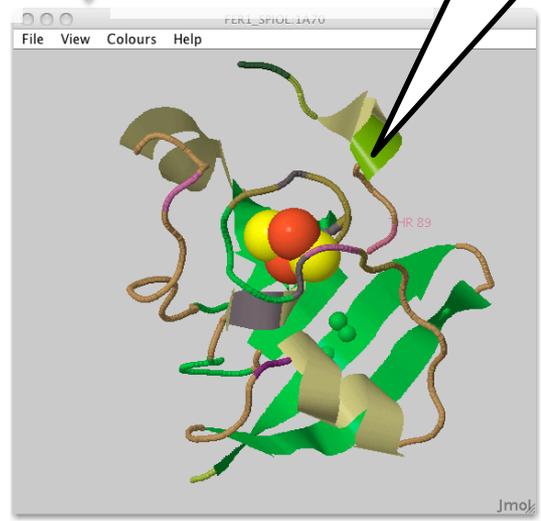
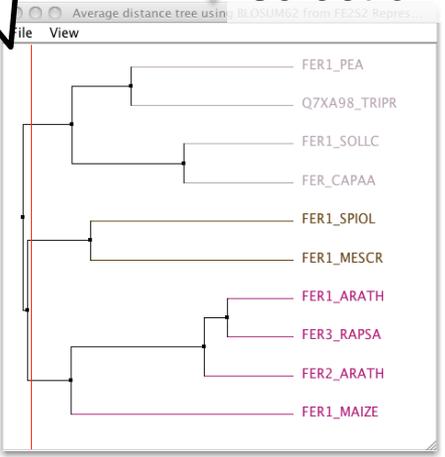


Linked tree viewer allows subgroups to be identified in alignment

Linked Jmol viewer shows one or more structures coloured by alignment



Group selections Colours and mouseovers



Associating structures with sequences

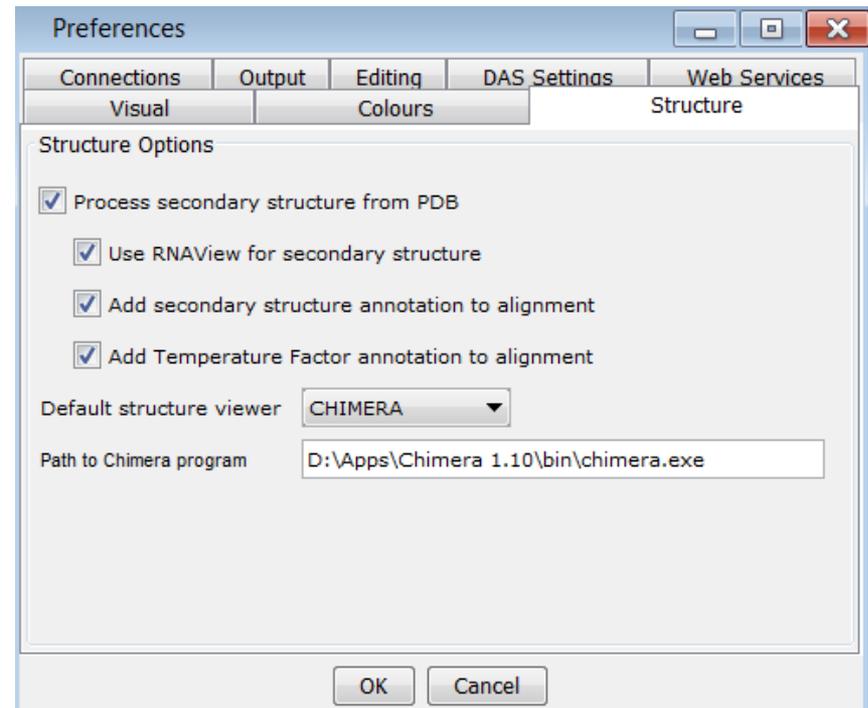
- Local PDB file
 - Attach PDB file to sequence manually
 - drag and drop to match files to sequences by ID
- Structures in the PDB database
 - Provide PDB id (and chain) for sequence
 - Discover references *via* sequence database



New Structure tab in Jalview Preferences

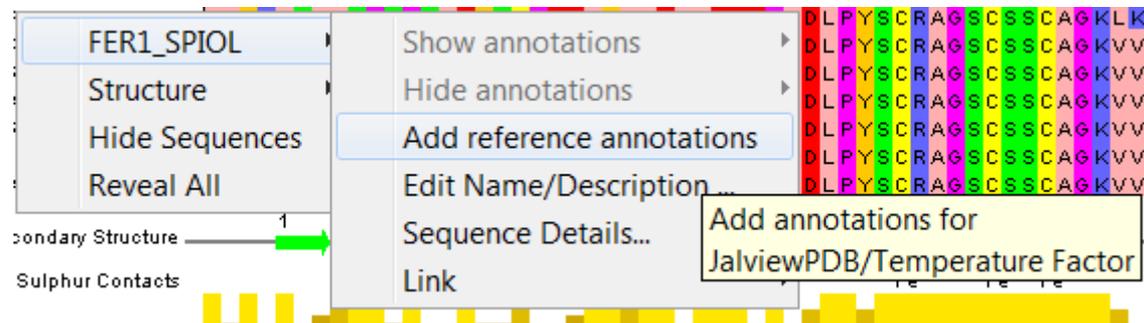
Checkboxes for:

- Secondary structure
 - Protein DSSP built in
 - RNAView web service
- Temperature factor
- Set default 3D viewer
 - Jmol
 - Chimera



When Protein annotation is enabled

- Add Reference Annotations
 - Available when any sequence associated annotation can be added to the current view
 - Access via Sequence ID and ‘Selection’ submenu



Protein Structures in Jalview

Sec 2.1. Exercise 14

- Task
 - Discover PDB structures for ferredoxin sequence(s)
 - **Note use**
Fetch Database Refs->UNIPROT->Uniprot
 - Save and load structures and manipulate colouring

Protein Structures in Jalview

Sec 2.1. Exercise 14

- Task
 - Discover PDB structures for ferredoxin sequence(s)
 - Save and load structures and manipulate colouring
- Questions
 - How does Jalview match up sequence data to structural data
 - Did anyone try ‘Add Reference Annotation’ ?

```
LIVNGGIIAPSFDDP.MDEKAREILQKLFPEHEVVMA.PGRELLL...L
YIANGGIIAPQFGDPIRDKEAIRVLSDTFPHHSVVGIENAREIVL.A...
LILNRRVFPVNGPASVDNDALNVYKTAMPGGYEIIGVKGASGTPWL.TDAL.R.HEVA
LITNKGVIIVPQYGDE.NDALALKQVQEMFPDREIVGV.NTVEVVY...
YVCNGGVVLCAFGDP.NDELAAGIFRRLFPERTVTLV.DARTIFA...G...
YTANGGIVFPLFNDP.MDEKAQEILQKLYPDRKIVGV.PAREILL...I...V...
```

Q8KCB6_CHLTE:1XKN Method: X-ray Chain:A...

File View Colours Jmol Help

Show Chain ▶

- All
- ✓ 1XKN:A
- ✓ 2CMU:A
- ✓ 1VKP:A
- ✓ 1VKP:B
- ✓ 2EWO:A
- 2EWO:B
- 2EWO:C
- 2EWO:D
- 2EWO:E
- 2EWO:F
- 2EWO:G
- 2EWO:H
- 2EWO:I
- 2EWO:J
- 2EWO:K
- 2EWO:L

CYS 366:A

Jmol

many structures can be shown in a single view

Structures can be Superimposed using the visible region of alignment

Superposing Structures using Alignments

Sec 2.1.4 – Exercise 15

- Task
 - Align structures using the ferredoxin alignment

If 'View all N structures' doesn't align structures:

- **Use Jmol->Align menu**
- Experiment with views to control what part of the alignment is used to superimpose the structures

Superposing Structures using Alignments

Sec 2.1.4 – Exercise 15

- Task
 - Align structures using the ferredoxin alignment
 - Experiment with views to control what part of the alignment is used to superimpose the structures
- Questions
 - What colourscheme would highlight the conserved parts of the structures ?
 - Which view gave the ‘best’ structure superposition ?
 - How did you decide this ?

Protein Disorder prediction

Exercise 27

- Complementary problem to secondary structure prediction
 - Recognise structured & unstructured domains
 - Predict holes in density maps (REM450)
 - Detect flexible loops ('HOTLOOPS')
- Programs provided by JABAWS 2 employ
 - Machine learning methods (**DisEMBL**)
 - Similarity to disordered sequences (**RONN**)
 - Empirical amino acid statistics (**IUPred**, **GlobPlot**)

Disorder Predictions from JABAWS

**JABAWS
Analysis
Service**

*Disorder
Predictions*

Create **annotation and
features**

<input checked="" type="checkbox"/> DisembiWS	<input checked="" type="checkbox"/> GlobPlotWS	<input checked="" type="checkbox"/> IUPredWS
Feature Type	Colour	Display
Protein Disorder		<input checked="" type="checkbox"/>
COILS		<input type="checkbox"/>
REM465		<input type="checkbox"/>
HOTLOOPS		<input type="checkbox"/>
Globular Domain		<input checked="" type="checkbox"/>

GlobPlotWS (Dydx)

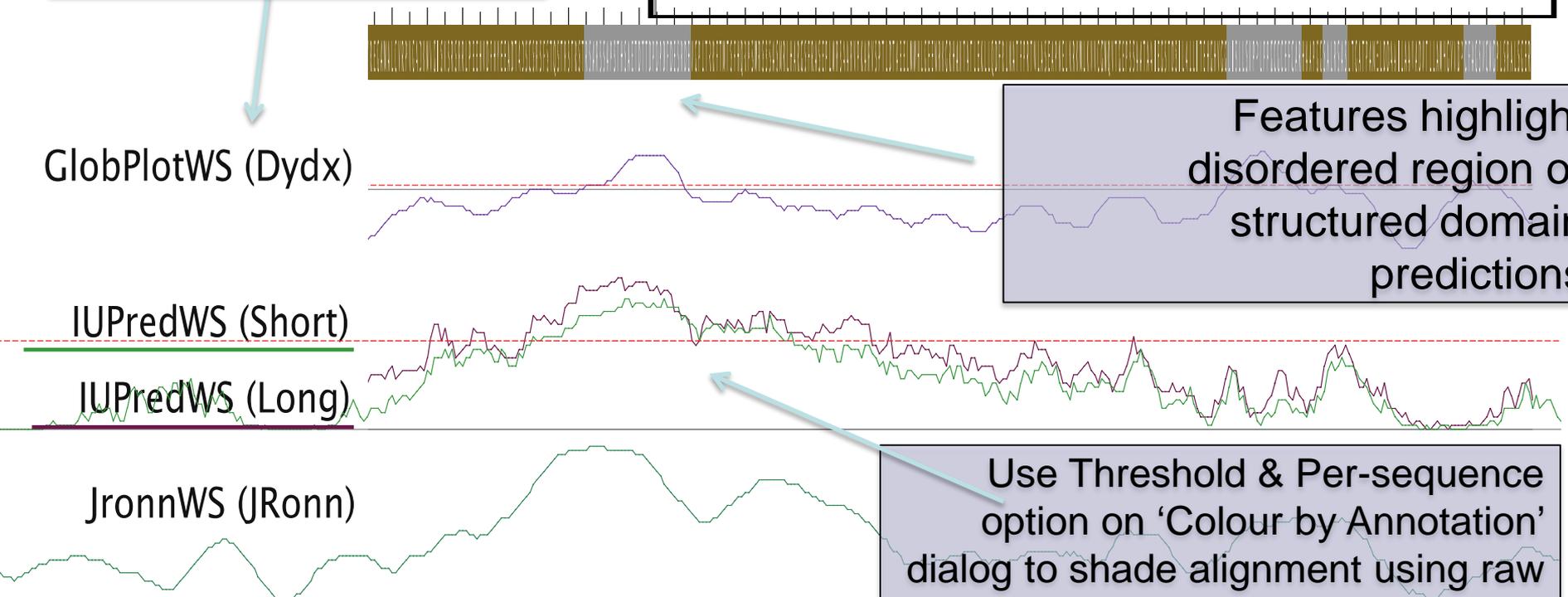
IUPredWS (Short)

IUPredWS (Long)

JronnWS (JRonn)

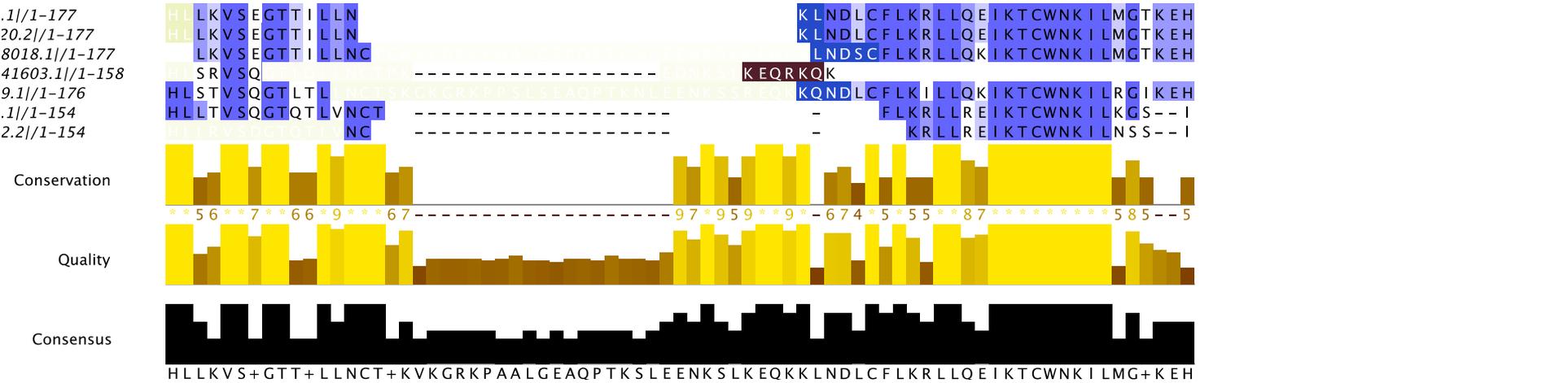
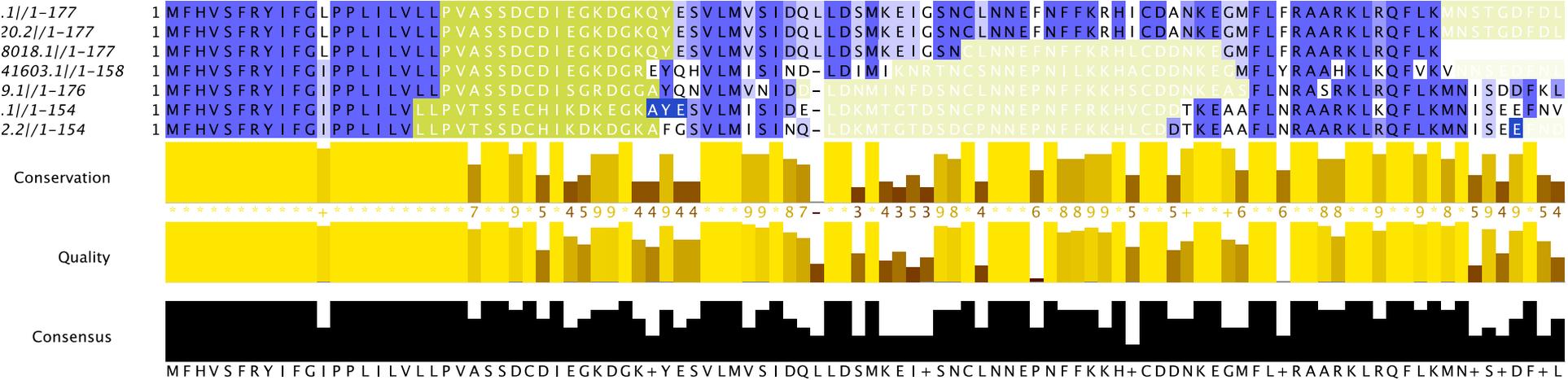
Features highlight
disordered region or
structured domain
predictions

Use Threshold & Per-sequence
option on 'Colour by Annotation'
dialog to shade alignment using raw
scores

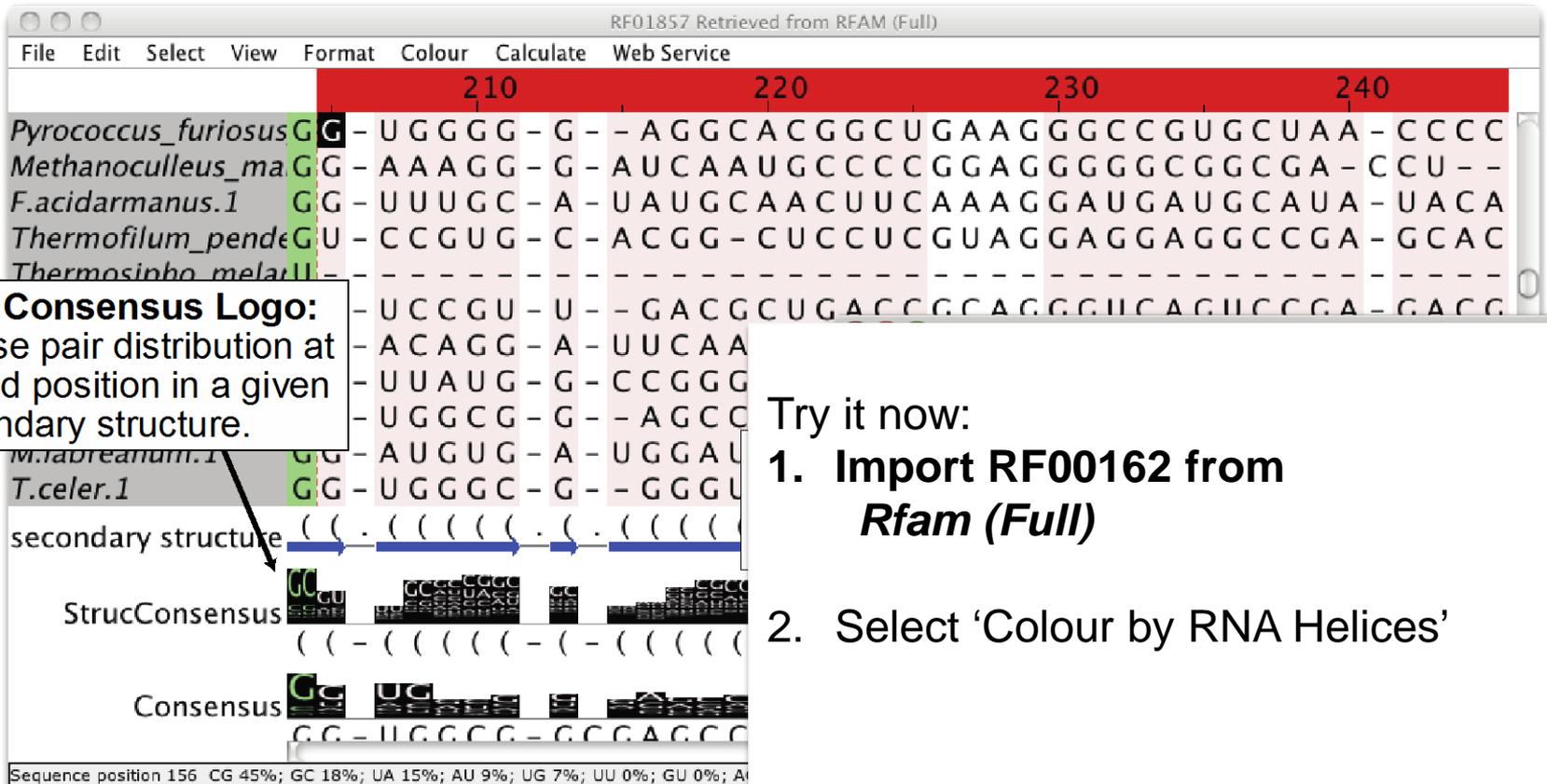


Disorder in Interleukin 7

~ ~ ~



RNA 2nd-ary Structure

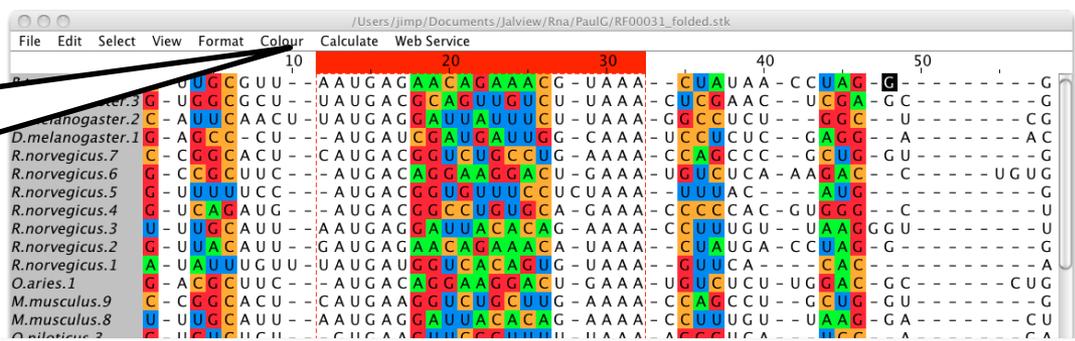


Structure Consensus Logo:
Shows base pair distribution at each paired position in a given RNA secondary structure.

- Try it now:
1. Import RF00162 from *Rfam (Full)*
 2. Select 'Colour by RNA Helices'



Selections and mouse positions shared between alignment view and VARNA



Try it now:

1. Open VARNA with
'Structure->View Structure->RNA Secondary Structure'
2. Explore difference between trimmed and untrimmed
3. Bonus points – add and link a **Jmol structure view**
Bacillus_amyloliquef.9 corresponds to PDB 3NPB

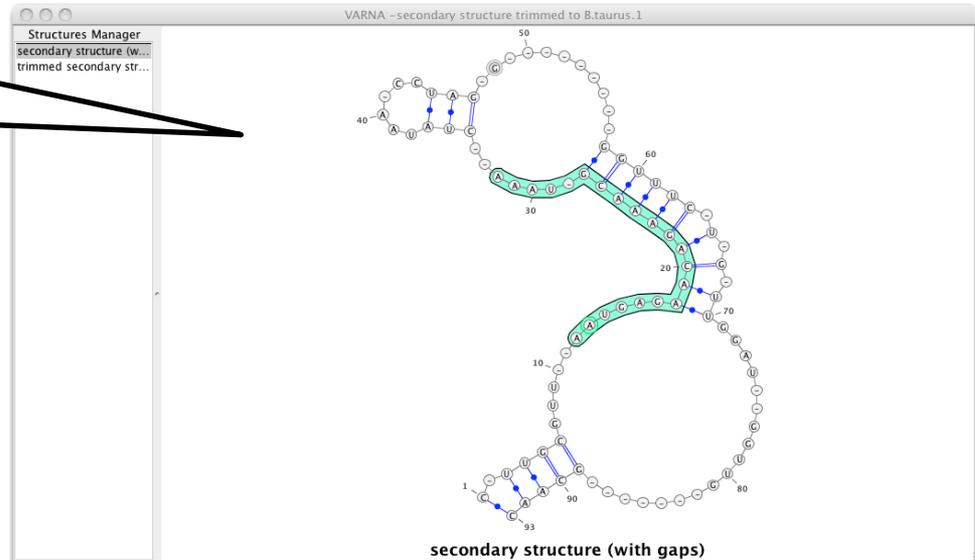
VARNA has a wide range of 2D RNA plots and supports interactive annotation



VARNA: Interactive drawing and editing of the RNA secondary structure

Kévin Darty, Alain Denise and Yann Ponty

Bioinformatics (2009) **25** 1974-1975



2.8.1 - Interactive Alignment based RNA Secondary Structure Prediction

ViennaRNA

Web Service

Alignment

Secondary Structure Prediction

Protein Disorder

Analysis

Conservation

Fetch DB References

JNet Secondary Structure Prediction

<http://www.compbio.dundee.ac.uk/jabaws>

RNAAliFold Prediction

Change RNAAliFold settings...

Modify settings for the RNAAliFold prediction. Use this to hide or show different results of the RNA calculation, and change RNA folding parameters

```
A-CCUUUGU--UAAGGGU-----UUG
A--CUAUGA-CCUAG-G-----GGU
A--GUUCA----CAC-----AGC
A-UGUCUCU-UGGAC-GC-----CUGGU
A-CCAGCCU--GCUG-GU-----GGG
A-CCUUUGU--UAAG-GA-----CUUG
A-AGGGUCA--UCC--A-----GAAA
```

2.8.1 - Interactive Alignment based RNA Secondary Structure Prediction

ViennaRNA

```

D.melanogaster.5/1-68 G - UGGCGCU
D.melanogaster.2/1-63 C - AUUCAAC
D.melanogaster.1/1-65 G - AGCC - CU
R.norvegicus.7/1-66 C - CGGCACU
R.norvegicus.6/1-67 G - CCGCUUC
R.norvegicus.5/1-62 G - UUUUUC
R.norvegicus.4/1-61 G - UCAGAUG
  
```

```

Consensus G UUUC
          UG ACGG
          G - UUUCAUU
  
```

RNAalifold Consensus B - H B K S W Y Y

Contact Probabilities

Centroid Structure

```

( . ( ( ( ( . . .
| | | | |
| | | | |
  
```

MFE Structure

```

( . ( ( ( ( . . .
| | | | |
| | | | |
  
```

StrucConsensus

```

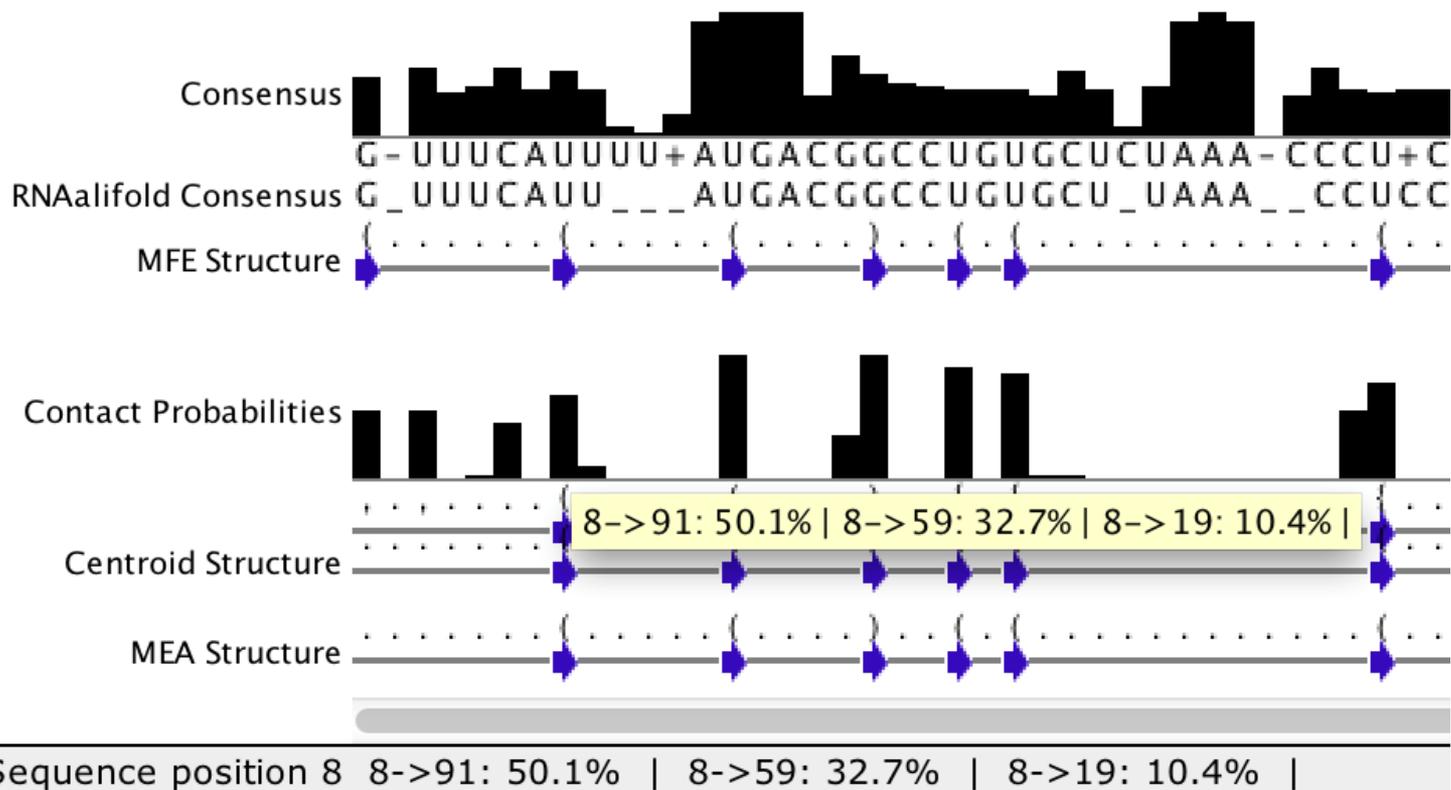
GC UACCUACC
CG UAGGCC
UA AGCCAUUA
UU UUUUUU
( - ( ( ( - - -
  
```

- Can be enabled for any view
- Updated if alignment changes
- settings & results saved in Jalview project

Implemented by our 2013 Summer student

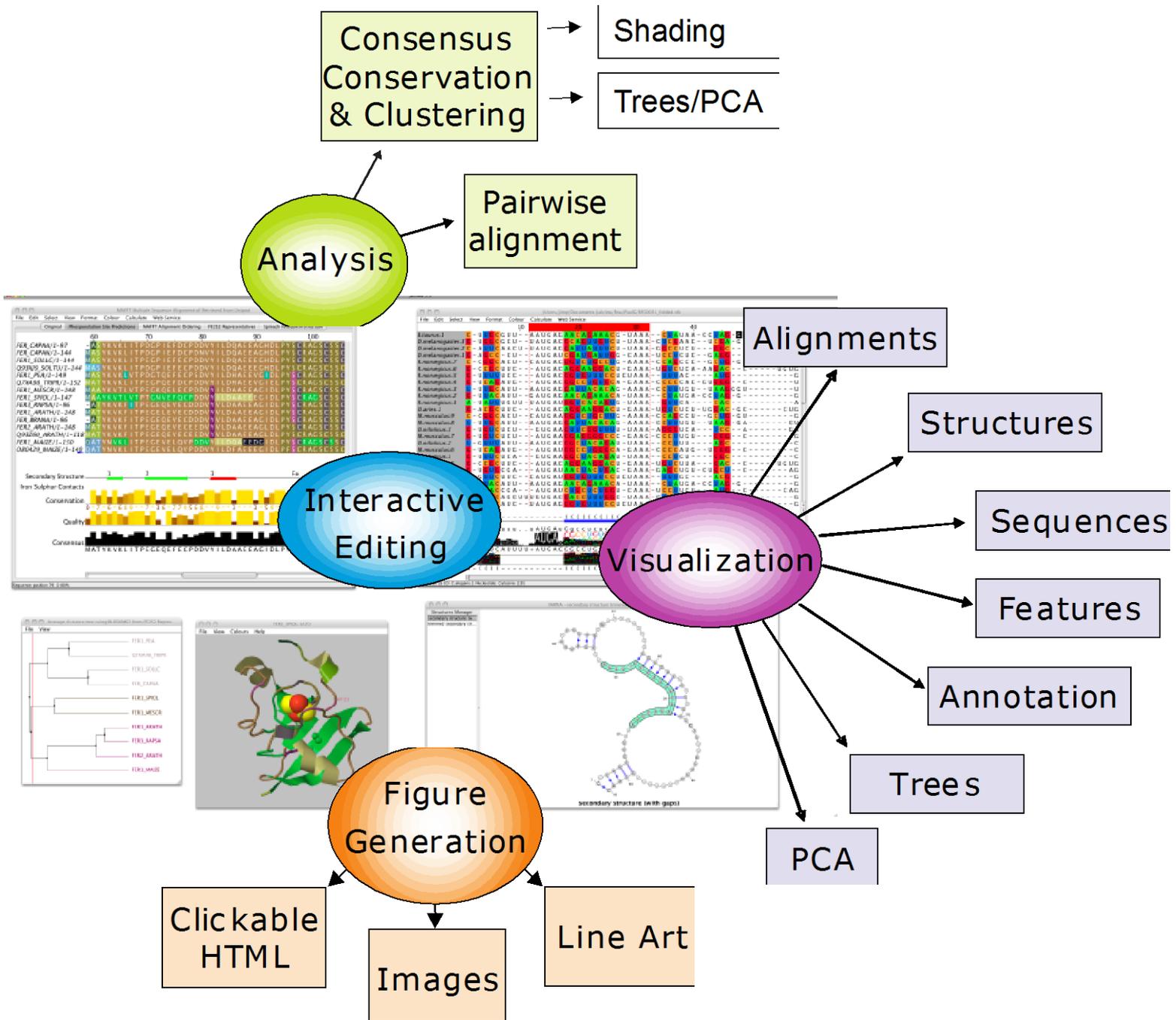
Tooltips show alternative base pairs

R.norvegicus.2/1-64 G - UUACA UU - - GAUGAGAA CAGAAACA - UAAA - - CUAUG
R.norvegicus.1/1-61 A - UAUUUGUU - UAUGAUGGUCACAGUG - UAAA - - GUUCA
O.aries.1/1-68 G - ACGCUUC - - - AUGACAGGAAGGACU - GAAA - UGUCUC



DNA and Protein in Jalview

- Discussed in Section 2.10 of manual
- From DNA to Protein
 - Calculations => Translate cDNA
 - View protein annotation on exons using EMBL records
- From protein to DNA
 - Recover DNA for proteins using EMBL cross references



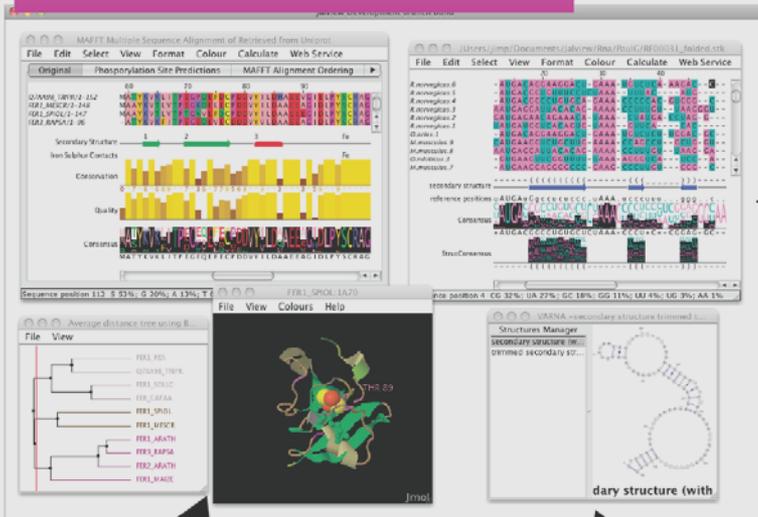
Jalview Flavours

Multi-windowed Web UI
Visualization & Analysis

Common Data & Analysis
Editing, messaging and
File Input/Output

Lightweight UI
Integrate with web sites

The Jalview Desktop



Figures

HTML
EPS
PNG



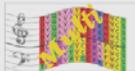
import
from web

Analysis Services

Protein 2ndary structure



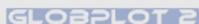
Alignment Analysis



Protein Disorder



Multiple alignment



Biological Data Services



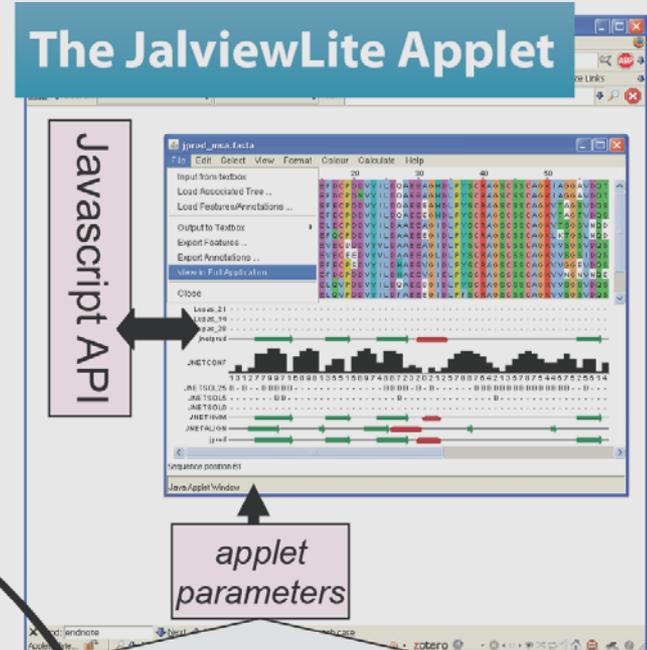
The JalviewLite Applet

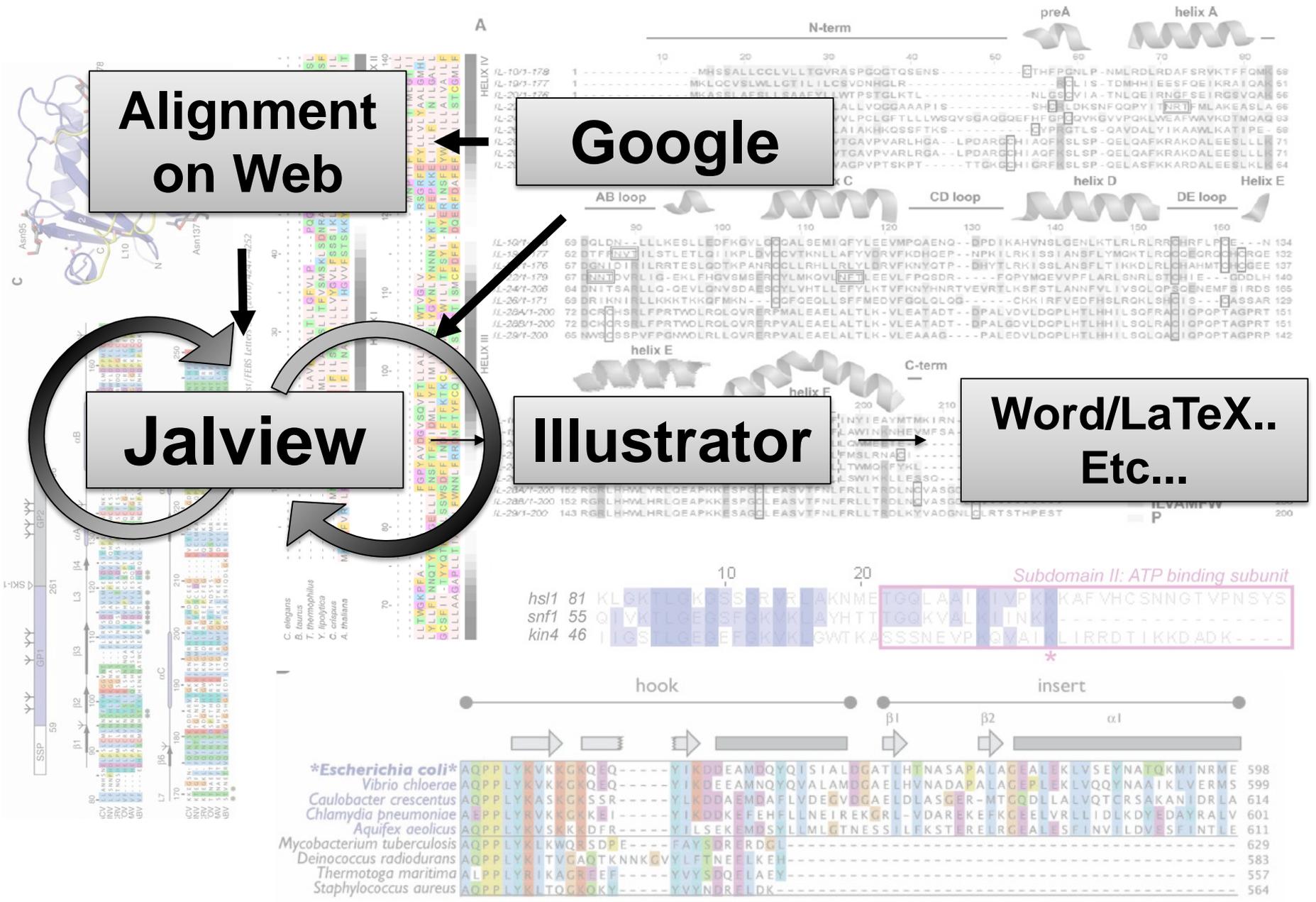
Javascript API

applet
parameters

Alignment
And
Annotation

Bioinformatics
Web Server





NEW FOR

2014

Jpred 3
Incorporating Jnet

ViennaRNA

Protein Disorder

UPred

RONN
Regional Order Neural Network

DisEMBL™

GLOBPLOT 2

Alignment Analysis

AACon

Multiple alignment

CLUSTAL T COFFEE

GLProbs MSAProbs

PROBCONS

Mafft MUSCLE W2
CLUSTAL

JABAWS 2.1

ViennaRNA

Ronny Lorenz et al. Algorithms for Molecular Biology, 2011 doi:10.1186/1748-7188-6-26

JPred3

Chris Cole, Jon Barber and Geoff Barton
NAR Web Server Issue 2008

doi:10.1093/nar/gkn238

Analysis

Interactive

Alignments

Structures

Sequences

MSAProbs: multiple alignment with pair HMMs

Yongchao Liu, Bertil Schmidt, and Douglas L. Maskell *Bioinformatics* 2010 26 1958 -1964

GLProbs – adaptive sequence alignment

Yongtao Ye et al. Proc. ACM BCB, 2013 pp.152-160

<http://sourceforge.net/projects/glprobs/>

DB

EUROPE

UniProt

ENA
European Nucleotide Archive

Distributed Annotation System

new
ures &

otation &

Protein Scores

Newick &
New
Hampshire
Extended

Things I haven't talked about ...

Select column by feature

ment of Retrieved from Uni

Calculate Web Service

MAFFT Align

80 90

DDVY I LDHAAEEVGI EL PY SC

DDVY I LDAAEEAGIDL PY SC

DDVY I LDAAEEEGIDL PY SC

DDVY VLDAEEAGIDL PY SC

DDVY VLDAEEAGIDL PY SC

DDVY VLDAEEAGIDL PY SC

EDVY VLDAEEAGLDL PY SC

EDVY VLDAEEAGLDL PY SC

DDVY I LDQAEEDGIDL PY SC

DDVY I LDFAEEEGIDL PY SC

3

6 8 * * 9 * * 3 * * * 3 * 5 9 * * 9 * *

Sequence Feature Settings

Feature Settings DAS Settings

uniprot netphos

1a70

Feature Type	Colour	Display
RESNUM		<input type="checkbox"/>
MOD_RES		<input checked="" type="checkbox"/>
VARIANT		<input checked="" type="checkbox"/>
METAL		<input checked="" type="checkbox"/>
PHOSPHORYLATION (S)		<input checked="" type="checkbox"/>
PHOSPHORYLATION (T)		<input checked="" type="checkbox"/>
PHOSPHORYLATION (Y)		<input checked="" type="checkbox"/>
CONFLICT		<input type="checkbox"/>
TURN		<input type="checkbox"/>
HELIX		<input type="checkbox"/>
STRAND		<input type="checkbox"/>

Sort by Score

Sort by Density

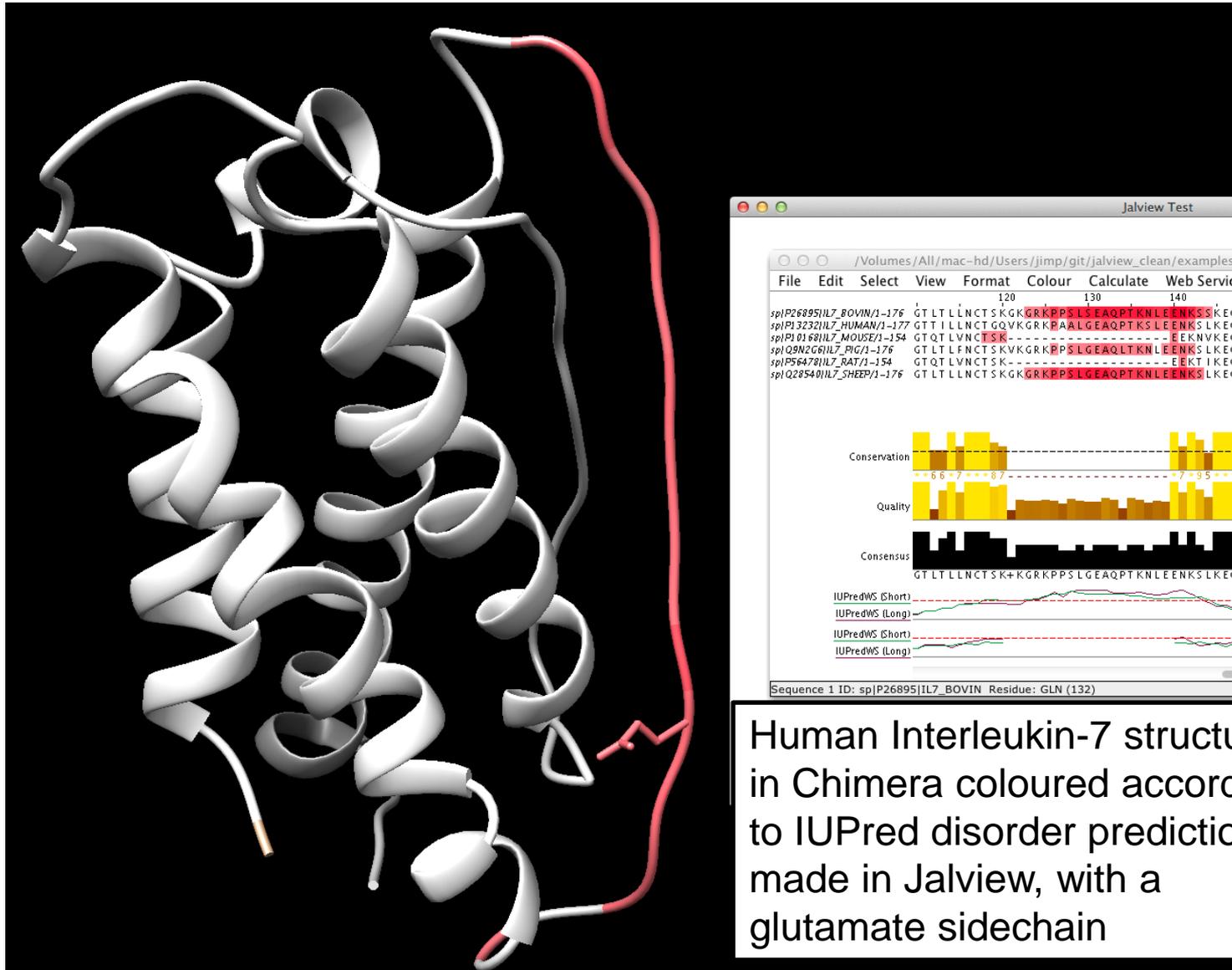
Graduated Colour

Select columns containing

Select columns that do not contain

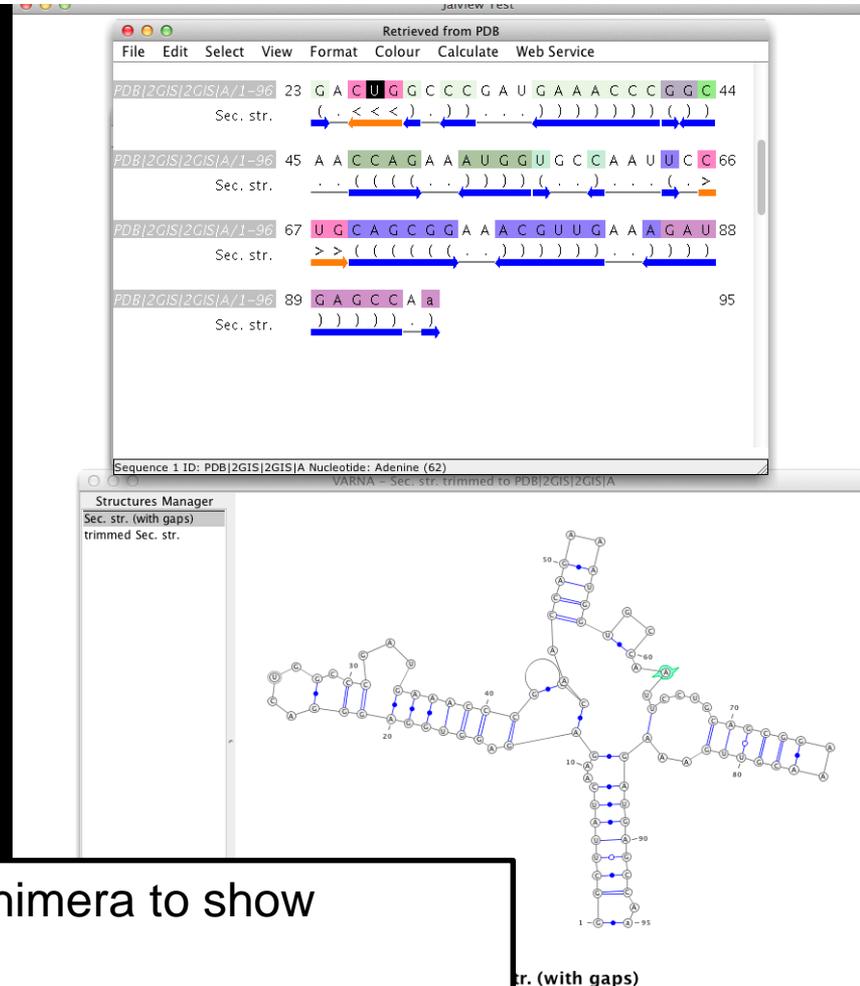
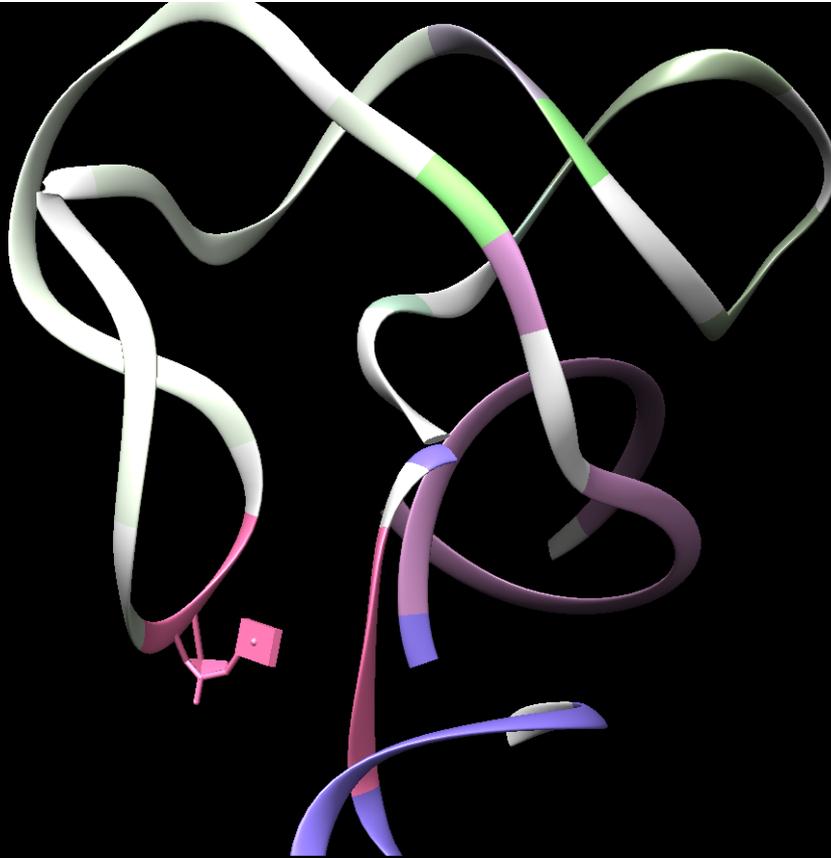
Invert Selection

Jalview and Chimera



Human Interleukin-7 structure in Chimera coloured according to IUPred disorder prediction made in Jalview, with a glutamate sidechain

Jalview + VARNA + Chimera



The screenshot shows the Jalview Desktop interface. The top window displays sequence alignment for PDB|2GIS|2GIS|A/1-96. The sequence is: 23 G A C U G G C C C G A U G A A A C C C G G C 44. Below the sequence, the secondary structure is shown with blue arrows and orange dots. The second window shows the sequence: 45 A A C C A G A A A U G G U G C C A A U U C G 66. The third window shows: 67 U G C A G C G G A A A C G U U G A A A G A U 88. The fourth window shows: 89 G A G C C A a 95. The bottom window shows the VARNA secondary structure prediction for the sequence, with a 3D ribbon diagram of the predicted structure.

Jalview Desktop uses UCSF Chimera to show structures:

- Optional - Enabled as a user preference
- Structures coloured & superposed like with Jmol

- Positional highlighting from Jalview->Chimera

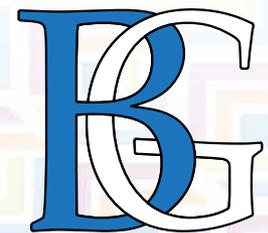
Jalview & Dundee Protein Resource Developers



Mungo
Carstairs

Tochukwu
(Charles)
Ofoegbu

Alexey
Drozdetskiy
Jnet Protein
prediction and
analysis services



supported by

wellcometrust





The Jalview developers
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Harvard & MIT.



James Cuff
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Andrew Waterhouse
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RNA Features

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Univ. Leipzig, Germany.

Yann Ponty (VARNA)
École Polytechnique,



T-COFFEE Scores
Paolo di Tomasso
Notredame Group, CRG, Spain.

Geoff Barton

- David Martin (**Teaching**)
- Sasha Sherstnev (**JABAWS**)
- Peter Troshin (**JABAWS**)
- Barry Strachan (**logo**)
- Tom Walsh (**Apache**)
- Ryan Maclaughlan (**CSS**)
- Andrew Millar (**Drupal**)
- All the Jalview users, and

...

JALVIEW



Jalview

www.jalview.org

supported by

