
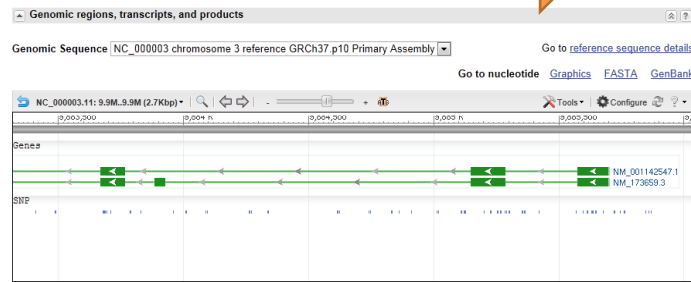



Genome browsers


Genome browsers

- Graphical interface for genomic data
- Shows information from biological databases mapped onto genomic sequence

Genomic coordinates 



Various annotations =
"tracks" 

NCBI Gene database  **Department of Informatics**
University of Oslo

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Structural Bioinformatics

UCSC Genome Browser

- Developed and maintained at the University of California, Santa Cruz (UCSC)
- Interactive website
- Access to genome sequence data from
 - Human genome
 - Latest assembly (GRCh37), but also earlier versions
 - Mouse, rat, and approx. 40 other mammals
 - Chicken, turkey, reptiles, frogs, and fish
 - Insects, nematodes, *S. cerevisiae* and more

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UCSC Genome Browser

D918-D923 *Nucleic Acids Research*, 2012, Vol. 40, Database issue
doi:10.1093/nar/gkr1055

Published online 15 November 2011

The UCSC Genome Browser database: extensions and updates 2011

Timothy R. Dreszer^{1,*}, Donna Karolchik^{1,*}, Ann S. Zweig¹, Angie S. Hinrichs¹, Brian J. Raney¹, Robert M. Kuhn¹, Laurence R. Meyer¹, Mathew Wong¹, Cricket A. Sloan¹, Kate R. Rosenbloom¹, Greg Roe¹, Brooke Rhead¹, Andy Pohl^{1,2}, Venkat S. Malladi¹, Chin H. Li¹, Katrina Learned¹, Vanessa Kirkup¹, Fan Hsu¹, Rachel A. Harte¹, Luvina Guruvadoo¹, Mary Goldman¹, Belinda M. Giardine³, Pauline A. Fujita¹, Mark Diekhans¹, Melissa S. Cline¹, Hiram Clawson¹, Galt P. Barber¹, David Haussler^{1,4} and W. James Kent¹

¹Center for Biomolecular Science and Engineering, School of Engineering, University of California Santa Cruz (UCSC), Santa Cruz, CA 95064, USA, ²Centre for Genomic Regulation (CRG), Barcelona, Spain, ³Center for Comparative Genomics and Bioinformatics, Huck Institutes of the Life Sciences, Pennsylvania State University, University Park, PA 16802 and ⁴Howard Hughes Medical Institute, UCSC, Santa Cruz, CA 95064, USA

Received September 15, 2011; Revised October 18, 2011; Accepted October 25, 2011

<http://genome.ucsc.edu>

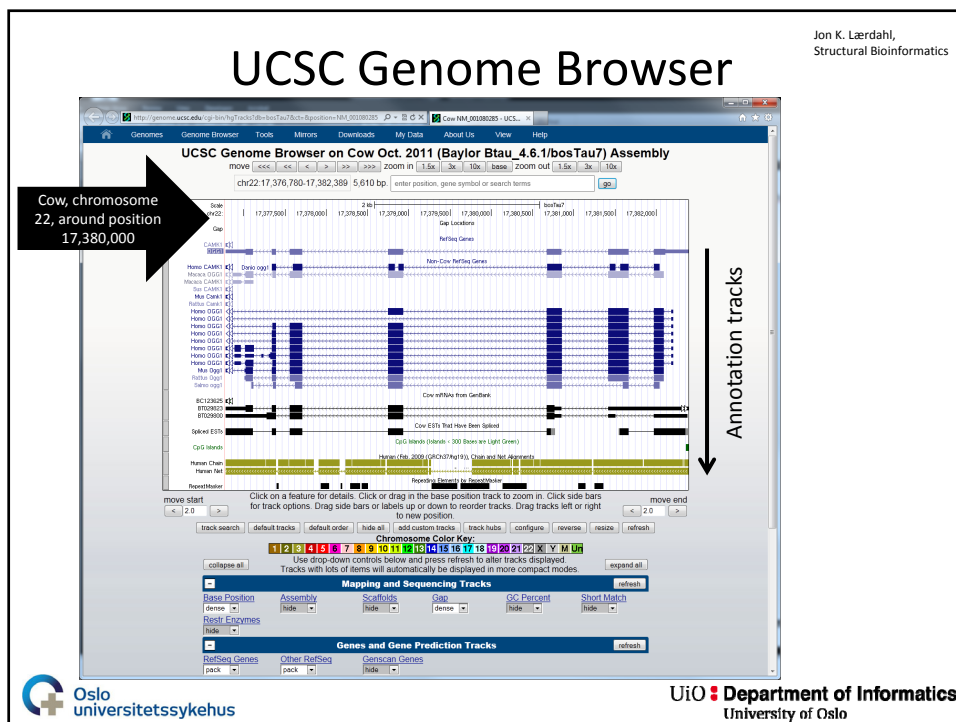
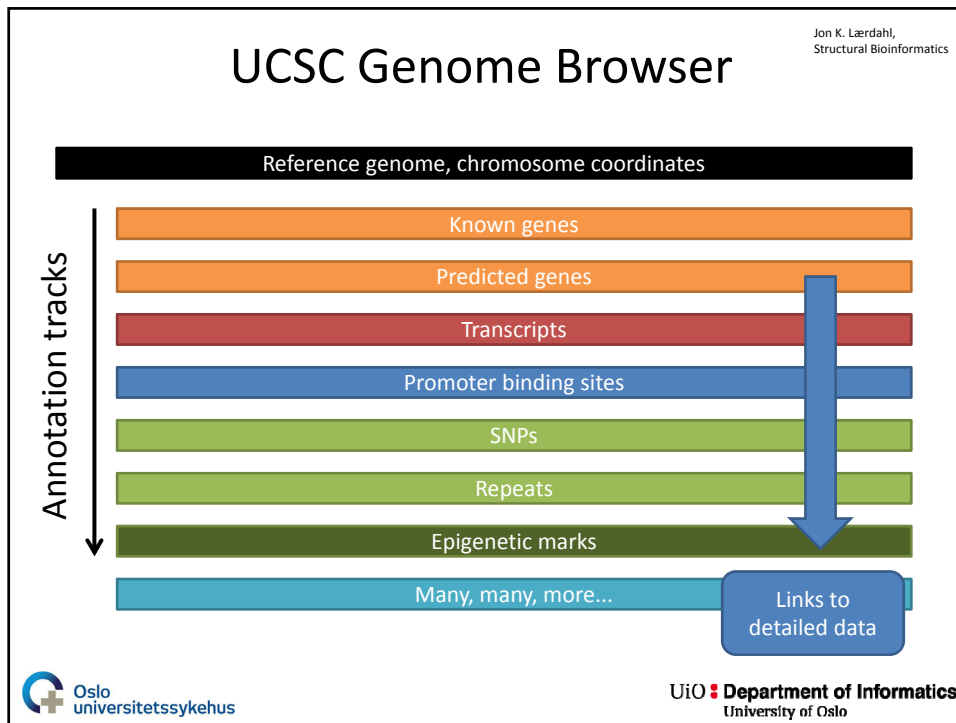
ABSTRACT

The University of California Santa Cruz Genome Browser (<http://genome.ucsc.edu>) offers online public access to a growing database of genomic sequence and annotations for a wide variety of organisms. The Browser is an integrated tool set for visualizing, comparing, analyzing and sharing both publicly available and user-generated genomic data sets. In the past year, the local database has been updated with four new species assemblies, and we anticipate another four will be released by the end

INTRODUCTION

The University of California Santa Cruz (UCSC) Genome Browser (1,2) at <http://genome.ucsc.edu> is a web-based set of tools providing access to a database of genome sequence and annotations for visualization, comparison and analysis by the scientific, medical and academic communities. Our primary mission is to provide timely and convenient open access to high-quality human genome sequence and annotations in a framework that enables easy exploration from genome-wide down to the base level. Annotation datasets, or 'tracks', on the human genome cover conservation and evolutionary compar-

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UCSC Genome Browser

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<http://genome.ucsc.edu>

Access to the databases and tools

Start here

General information

News, updates, announcements

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UCSC Genome Browser

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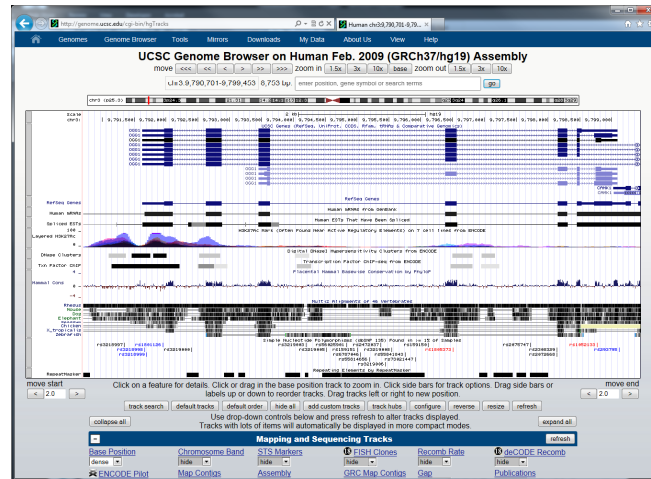
Examples of searching options – correct query format

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UCSC Genome Browser brief demo

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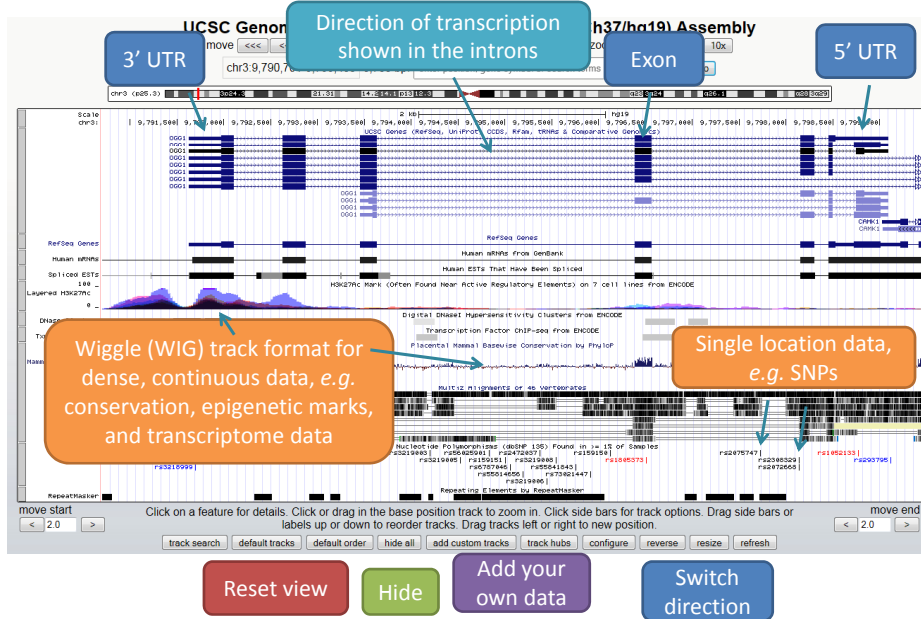


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Different kinds of data

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ENCODE data in UCSC

Jon K. Lærdahl,
Structural Bioinformatics

Published online 30 October 2010

Nucleic Acids Research, 2011, Vol. 39, Database issue D871–D875
doi:10.1093/nar/gkq1017

ENCODE whole-genome data in the UCSC genome browser (2011 update)

Brian J. Raney^{1*}, Melissa S. Cline¹, Kate R. Rosenbloom¹, Timothy R. Dreszer¹, Katrina Learned¹, Galt P. Barber¹, Laurence R. Meyer¹, Cricket A. Sloan¹, Venkat S. Malladi¹, Krishna M. Roskin¹, Bernard B. Suh¹, Angie S. Hinrichs¹, Hiram Clawson¹, Ann S. Zweig¹, Vanessa Kirkup¹, Pauline A. Fujita¹, Brooke Rhead¹, Kayla E. Smith¹, Andy Pohl¹, Robert M. Kuhn¹, Donna Karolchik¹, David Haussler^{1,2} and W. James Kent¹

¹Center for Biomolecular Science and Engineering, School of Engineering and ²Howard Hughes Medical Institute, University of California Santa Cruz (UCSC), Santa Cruz, CA 95064, USA

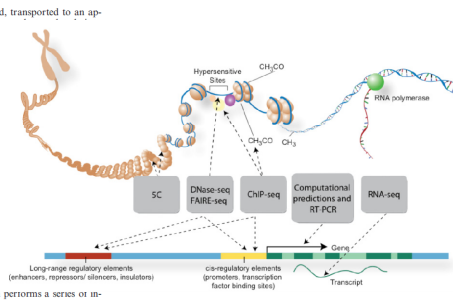
Received September 15, 2010; Accepted October 9, 2010

ABSTRACT

The ENCODE project is an international consortium with a goal of cataloguing all the functional elements in the human genome. The ENCODE Data Coordination Center (DCC) at the University of California, Santa Cruz serves as the central repository for ENCODE data. In this role, the DCC offers a collection of high-throughput, genome-wide data generated with technologies such as ChIP-Seq, RNA-Seq, DNA digestion and others. This data helps illuminate transcription factor-binding sites, histone marks, chromatin accessibility, DNA methylation, RNA expression, RNA binding and other cell-state indicators. It includes sequences with quality scores, alignments, signals calculated from the alignments, and in most cases, element or peak calls calculated from the signal data. Each data set is available for visualization and download via the UCSC Genome Browser (<http://genome.ucsc.edu>). ENCODE data can also be retrieved using a metadata system that captures the experimental parameters of each assay. The ENCODE web portal at UCSC (<http://encodeproject.org/>) provides information about the ENCODE data and links for access.

into RNA, which might be spliced, transported to an appropriate cellular compartment. This process is regulated by transcription factors to the DNA factors to the RNA and RNA traits are determined as differences in gene regulation as differences in gene expression. The goal of the ENCODE project is to catalog all the functional elements in the human genome. This process is regulated by DNA methylation, chromatin accessibility, transcription factors to the DNA factors to the RNA and RNA traits are determined as differences in gene regulation as differences in gene expression. The role of the ENCODE Data Coordination Center (DCC) is to organize and display data from the consortium, and to ensure specific quality standards when it receives data from any lab. Before a lab submits any data, it must agree to a data agreement that defines the data and associated metadata. The DCC data to ensure consistency with the data onto a test server, and coordinates with the data into a consistent set of tracks. With the DCC Quality Assurance team performs a series of in-

<http://genome.ucsc.edu/ENCODE/aboutScaleup.html>



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Ensembl Genome Browser

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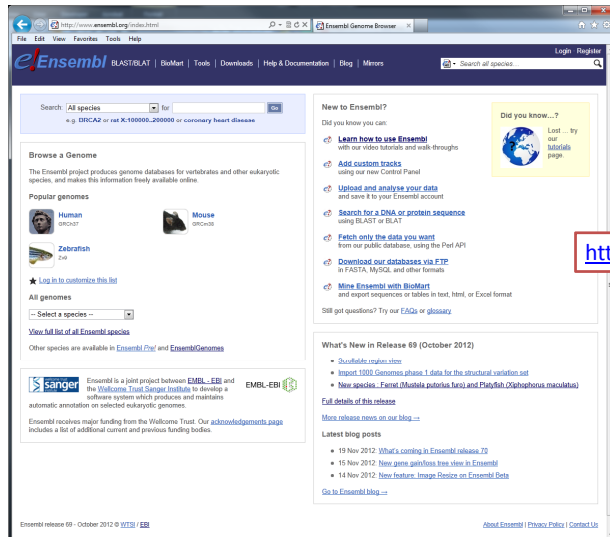
- Joint project between EMBL-EBI and the Wellcome Trust Sanger Institute
- Central resource for studying genomes of vertebrates
 - Mainly chordates, but some few extra (*e.g. C. elegans* and *S. cerevisiae*)
 - Updated several times a year with new genome assemblies and new species
 - Annotations of genomes (*e.g.* genes and their splice variant, SNPs) added by the Ensembl pipeline
 - Automatic gene prediction (with or without experimental evidence) & some curator input

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Ensembl Genome Browser



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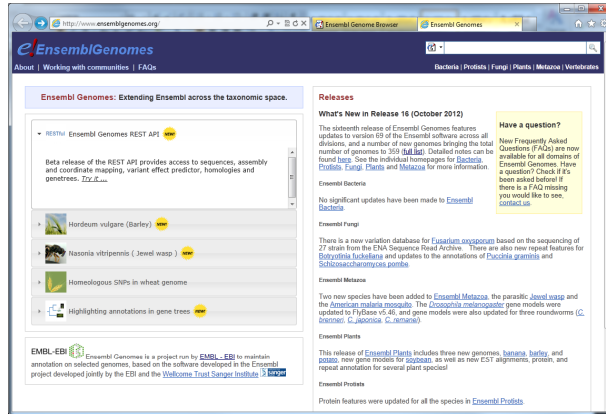


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EnsemblGenomes

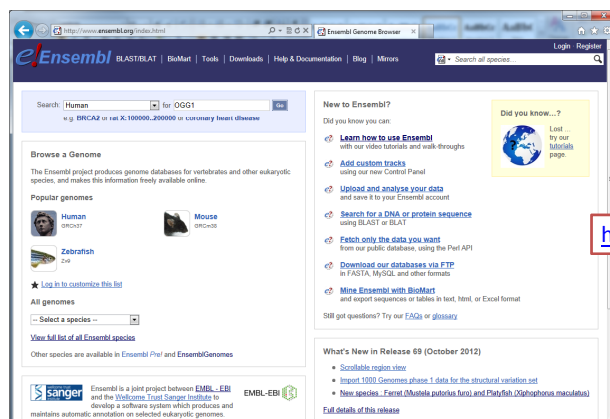
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- Bacteria, protists, fungi, plants and other metazoa (359 genomes)

Ensembl Genome Browser

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Explore!

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Now something different!



Not a genome browser!

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- Galaxy is a platform (open, web-based) for computational medical projects and bioinformatics
 - Accessible: Not necessary to know programming, Unix, or how to install programs
 - Reproducible: You can build and store complete workflows, pipelines, and the full computational analysis
 - Transparent: Users can publish and share whole workflows
- A bioinformatics workflow management system

Reproducibility



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Repeatability of published microarray gene expression analyses

John P A Ioannidis¹⁻³, David B Allison⁴, Catherine A Ball⁵, Issa Coulibaly⁴, Xiangqin Cui⁴, Aedin C Culhane^{6,7}, Mario Falchi^{8,9}, Cesare Furlanello¹⁰, Laurence Game¹¹, Giuseppe Jurman¹⁰, Jon Mangion¹¹, Tapan Mehta⁴, Michael Nitzberg⁵, Grier P Page^{4,12}, Enrico Petretto^{11,13} & Vera van Noort¹⁴



Given the complexity of microarray-based gene expression studies, guidelines encourage transparent design and public data availability. Several journals require public data deposition and several public databases exist. However, not all data are publicly available, and even when available, it is unknown whether the published results are reproducible by independent scientists. Here we evaluated the replication of data analyses in 18 articles on microarray-based gene expression profiling published in *Nature Genetics* in 2005–2006. One table or figure from each article was independently evaluated by two teams of analysts. We reproduced two analyses in principle and six partially or with some discrepancies; ten could not be reproduced. The main reason for failure to reproduce was data unavailability, and discrepancies were mostly due to incomplete data annotation or specification of data processing and analysis. Repeatability of published microarray studies is apparently limited. More strict publication rules enforcing public data availability and explicit description of data processing and analysis should be considered.

Figure 1 Summary of the efforts to replicate the published analyses.

Reproducibility

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UNMATCHED MESSAGES

The swapping out of a single base in a section of messenger RNA alters the genetic information transcribed from the DNA before it is translated into a protein.

Transcription: A complementary strand of RNA is produced.

Translation: A ribosome reads this codon. A three-base sequence codes for a specific amino acid. Relevant amino acid is added. Growing polypeptide chain. Final protein.

RNA editing: A single base is altered. A different amino acid is added. A different protein is produced.

432 | NATURE | VOL 473 | 26 MAY 2011 © 2011 Macmillan Publishers Limited. All rights reserved.

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NEWS & COMMENT > News > 2012 > November > Article

NATURE | NEWS

RNA editing study under intense scrutiny

Debate highlights pitfalls in interpreting genomic data.

Erika Check Hayden
15 March 2012

Three groups¹⁻³ today throw fuel on the debate surrounding a researcher's claim to have discovered a twist in the mechanism whereby genes are translated into proteins. The paper, published in *Science* last May⁴, suggested a revision in the central dogma, which holds that the RNA transcripts used as templates for protein assembly are generally faithful matches to the original DNA. In technical commentaries published today in *Science*, the groups suggest that errors in multiple aspects of the study, led by Vishal Cheung of the University of Pennsylvania in Philadelphia, seriously undermine his claim to have found a new mechanism of genetic regulation.

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1. Quantum cryptography conquers noise problem
Nature | 19 November 2012
2. Great apes go through mid-life crisis
Nature | 19 November 2012

Li, M. *et al.* *Science* **33**, 53 (2011), July 2011:

- >10,000 sites in the human genome where an RNA sequence did not match the sequence of the DNA
- Evidence for a new mechanism of RNA editing?
- Central dogma is wrong in 10,000 places?

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- Galaxy is a platform (open, web-based) for computational medical projects and bioinformatics
 - Accessible: Not necessary to know programming, Unix, or how to install programs
 - **Reproducible: You can build and store complete workflows, pipelines, and the full computational analysis**
 - **Transparent: Users can publish and share whole workflows**
- By the way, Galaxy is written in Python...
- Developed by the labs of Anton Nekrutenko (Penn State University) and James Taylor (Emory University)

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30 June - 2 July 2013
UiO University of Oslo



- Biportal -> Galaxy
- Galaxy at UiO submitting jobs to the Abel cluster

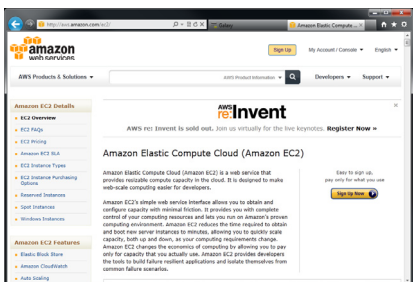



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- Can be run on a free public server at Penn State
- You can install Galaxy on your own server or computer cluster (soon on Abel)
- You can run Galaxy in the cloud

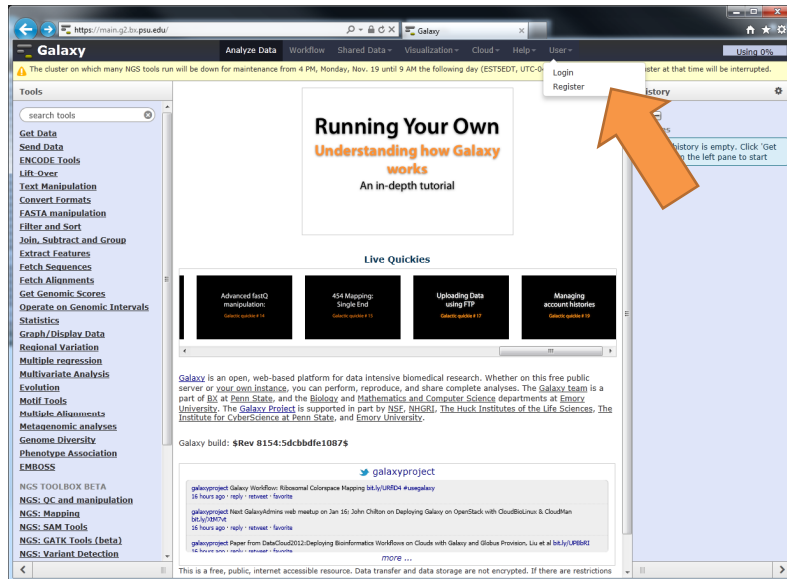




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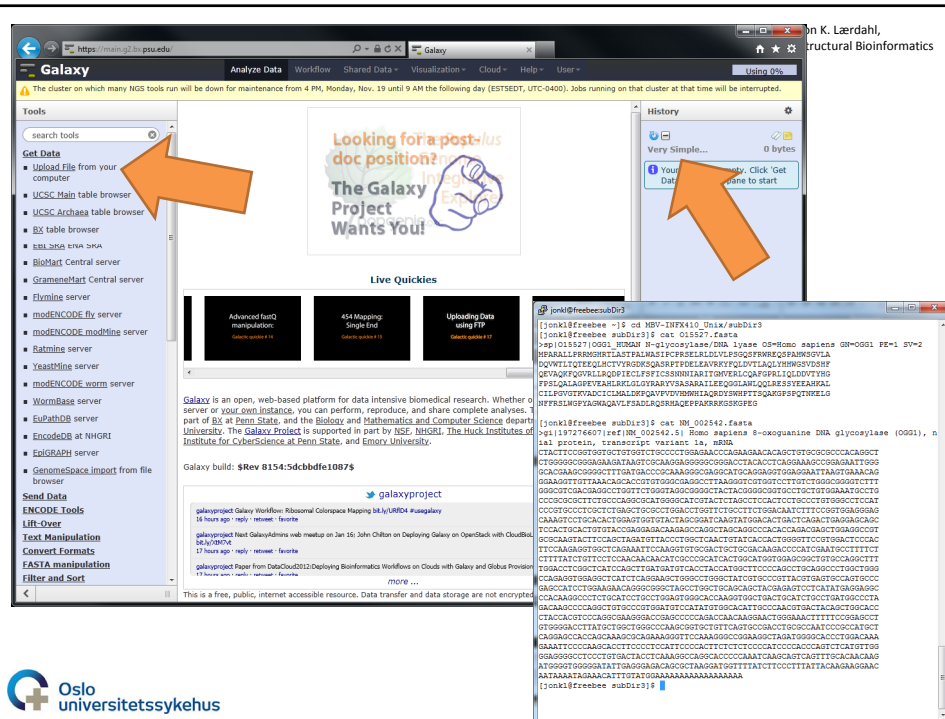
Free public server:

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The screenshot shows the Galaxy web interface. The 'Upload File (version 1.1.3)' dialog is open, with the 'File Format' set to 'fasta'. An orange arrow points to the 'Execute' button. Another orange arrow points to the 'Files uploaded via FTP' table, which is currently empty. A terminal window in the foreground shows the following commands and output:

```
jonk@freebees01r:~$ cat NM_002545.fasta
[jonk@freebees01r:~]$ cd NM_002545
[jonk@freebees01r:~]$ cat 01527.fasta
>>>[15527] OGG1_HUMAN H-glycosylase/DNA lyase OS=Momo sapiens GN=OGGI PE=1 SV=2
MPAALLFPNGSRTLSTALMNSTPCPPSELRLDVLVPSQSPFPAKQSPAKNSQVLA
DQVTTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTT
QVVAQFQVRLRDPQFIECLFSTCSNNHILRTYMPFERCQAFQFPLQLDQVTRG
FFGQALAEFEVEMRSHGLQFNPAVYKASALILEGQQALMGQLRSEFHEENRAL
CILPQVTFVADICLMLKQFQVFDVVRHRIAGQDYRSHFTTSQMGPSFQNKELG
MFRFSLMGVAGAAVLFASALRQSRGAEQEPAPRBRSGS92FG
```



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This screenshot shows the Galaxy interface after a job has been submitted. A green notification box states: "The following job has been successfully added to the queue: 1: NM_002545.fasta". An orange arrow points to the 'History' tab, which now displays the job details:

```
1: NM_002545.fasta
1 1 1 1
Very Simple... 0 bytes
Format: fasta, database: 2
Info: uploaded fasta file
[0] [0]
```

A second orange arrow points to the 'Tools' pane on the left. A blue box with the text "After a while..." is positioned between the two screenshots, indicating the passage of time.



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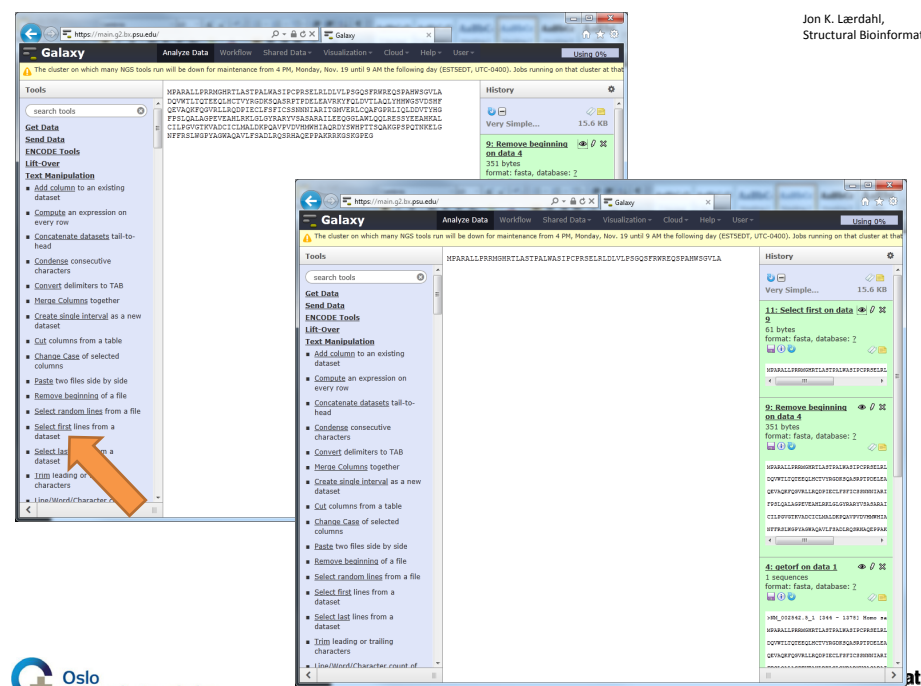
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The screenshot shows the Galaxy web interface. The main panel displays a workflow with a 'getorf' tool. The history panel on the right shows a list of jobs. An orange arrow points to the job '4: getorf on data 1'. A text box next to the arrow says 'Where is 3? I did a mistake and had to remove it'. The left sidebar shows a list of tools, including 'standem', 'extractfeat', 'extractreg', 'freq', 'fuzmex', 'fuztran', 'sarnier', 'anagec', 'getorf', 'haldutunhalla', 'hmoment', 'isp', 'infogan', 'isochore', 'linda', and 'margin'. The top navigation bar includes 'Galaxy', 'Analyze Data', 'Workflow', 'Shared Data', 'Visualization', 'Cloud', 'Help', and 'User'. A warning message at the top states: 'The cluster on which many NGS tools run will be down for maintenance from 4 PM, Monday, Nov. 19 until 9 AM the following day (ESTSEDT, UTC-0400). Jobs running on that cluster at that time will be interrupted.'

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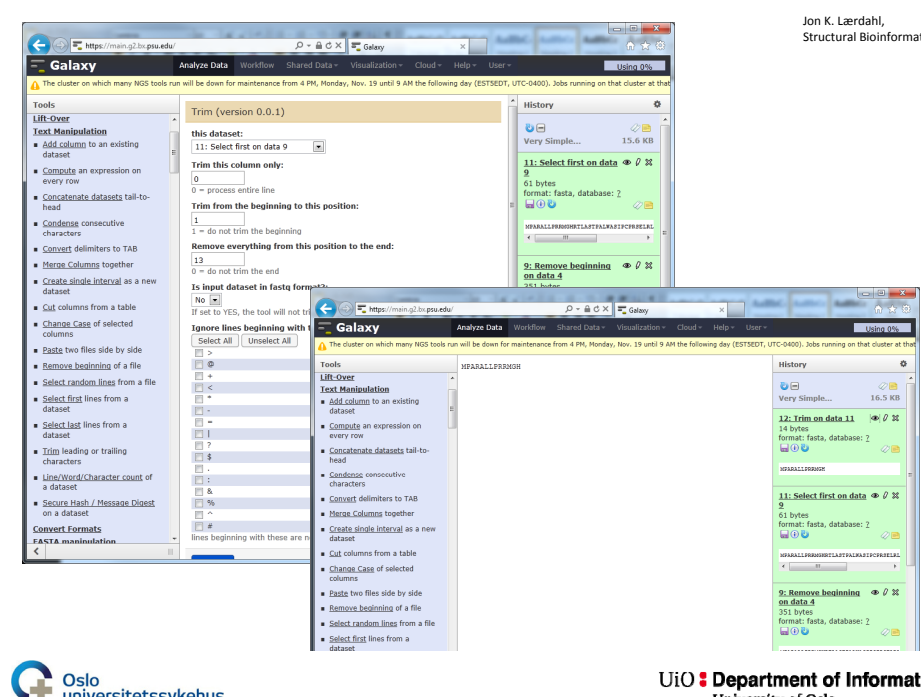
The screenshot shows the Galaxy web interface with the 'Remove beginning (version 1.0.0)' tool configuration. The left sidebar shows a list of tools, including 'Remove beginning of a file'. An orange arrow points to this option. The main panel shows the tool configuration with 'Remove first:' set to '1' and 'from:' set to '4: getorf on data 1'. The 'What it does' section states: 'This tool removes a specified number of lines from the beginning of a dataset.' The 'Example' section shows input and output files. The history panel on the right shows a list of jobs. The top navigation bar includes 'Galaxy', 'Analyze Data', 'Workflow', 'Shared Data', 'Visualization', 'Cloud', 'Help', and 'User'. A warning message at the top states: 'The cluster on which many NGS tools run will be down for maintenance from 4 PM, Monday, Nov. 19 until 9 AM the following day (ESTSEDT, UTC-0400). Jobs running on that cluster at that time will be interrupted.'



The screenshot shows the Galaxy web interface. On the left, the 'Tools' panel is open, displaying a list of tools under 'Text Manipulation'. An orange arrow points to the 'Select first lines from a dataset' tool. The main workspace shows a workflow with a 'Very Simple...' tool. The 'History' panel on the right shows a list of jobs, including '9: Remove beginning on data_4' and '11: Select first on data_4'. A warning banner at the top indicates that the cluster will be down for maintenance from 4 PM on Monday, Nov. 19 until 9 AM the following day.

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The screenshot shows the Galaxy web interface with the 'Trim (version 0.0.1)' tool configuration panel open. The tool is set to 'Trim this column only' with '1' selected. The 'Remove everything from this position to the end' is set to '13'. The 'History' panel on the right shows a list of jobs, including '11: Select first on data_4' and '12: Trim on data_11'. The 'Tools' panel on the left is also visible, showing various text manipulation tools.

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Workflow Canvas | Workflow constructed from history 'Very Simple...'

- Create a "workflow" from your history
- Edit workflow
 - Edit settings for all "modules"
 - Add new modules
- Share workflow
 - Share on a website?
 - As supplementary material for publication?

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- A *very* simple demonstration
- Galaxy can quite easily answer questions like:
 - Which exon in the human genome contains the largest number of SNPs?
 - How many exons are there on mouse chromosome 1?
- Very good for making pipeline to analyze HTS data
 - CHIP-seq, RNA-seq etc
- If you are doing this kind of work, Galaxy might be something to consider!
- Try it yourself? Check out the Galaxy 101 screencast?