ENCODE Standards, Data and Access

Advanced workshop on integrative analysis using ENCODE and Roadmap Epigenomics data

J. Michael Cherry Stanford University, Department of Genetics October 10, 2015





Workshop outline

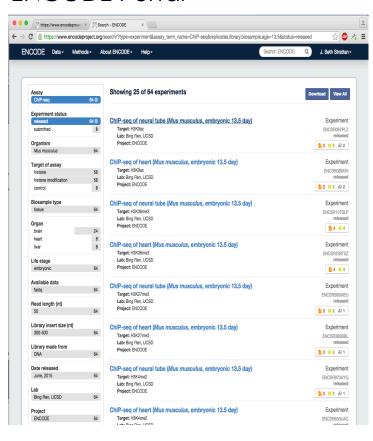
<u>Data Access & processing</u>

Interpretation

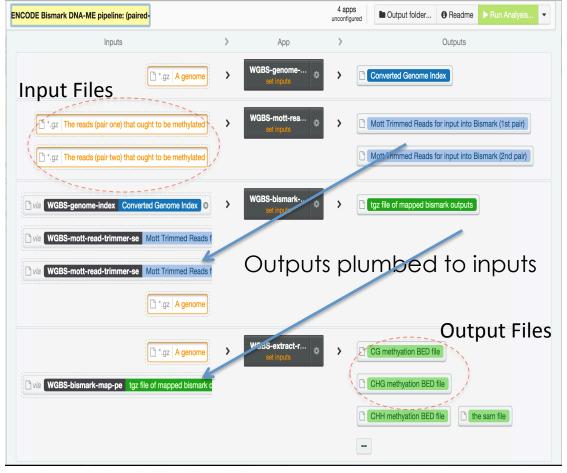
Visualization

Advanced Analysis

ENCODE Portal



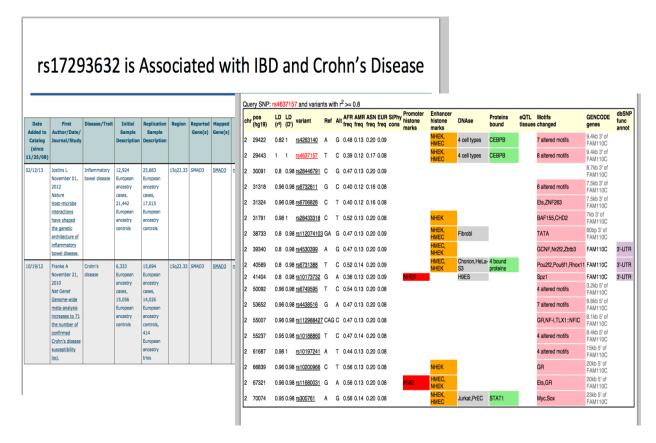
ENCODE Processing Pipelines



Workshop outline

Data Access & processing -> Interpretation -> Visualization -> Advanced Analysis

Jill Moore: HaploReg and RegulomeDB



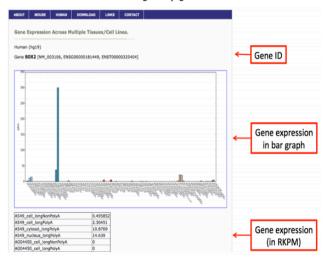
Workshop outline

Data Access & processing -> Interpretation -> <u>Visualization</u> -> <u>Advanced Analysis</u>

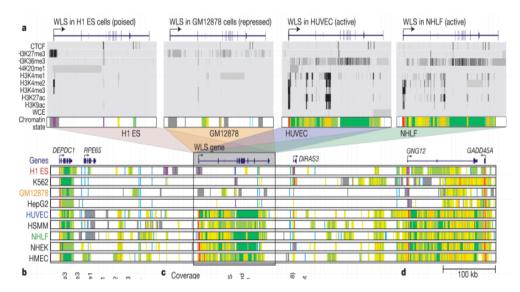
Yanli: Element and 3D Browser



You will be re-directed to the following result page.



Jason Ernst: Genome annotation ChromHMM



Goals for this section

- Equip you with the knowledge and tools to explore ENCODE on your own
 - Learn how to find and download data
 - Learn about the features of the portal
 - Learn about the rich metadata available that describes and contextualizes data
 - Learn about the ENCODE uniform processing pipelines

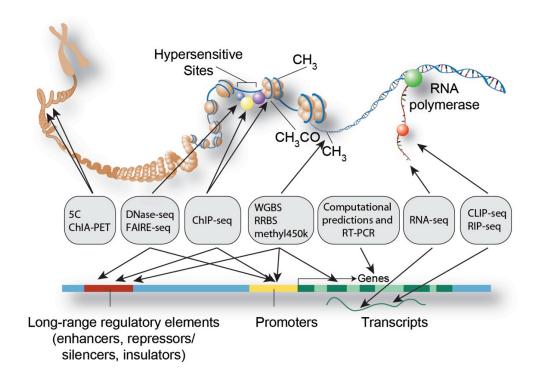
ENCODE has produced a lot of data

ENCODE

From 2003 to present >5000 experiments >3500 biosamples Hundreds of terabytes of files

Roadmap Epigenomics

>3127 experiments (ChIP-seq, Dnase-seq, RNA-seq and bisulfite-seq >1200 biosamples





FEATURES



Reproducibility crisis: Blame it on the antibodies

Antibodies are the workhorses of biological experiments, but they are littering the field with false findings. A few evangelists are pushing for change.

Nature (19 May 2015)

antibodies



Statistical errors

P values, the 'gold standard' of statistical validity, are not as reliable as many scientists assume. *Nature* (12 February 2014) quality control & measures of confidence



Replication studies: Bad copy

In the wake of high-profile controversies, psychologists are facing up to problems with replication. *Nature* (16 May 2012)

replication in experimental design

NEWS AND ANALYSIS



Irreproducible biology research costs put at \$28 billion per year

Study calculates cost of flawed biomedical research in the United States. *Nature* (09 June 2015)



Sluggish data sharing hampers the reproducibility effort

Initiative trying to validate 50 cancer papers finds difficulty in accessing original study data. *Nature* (03 June 2015)

data not readily accessible

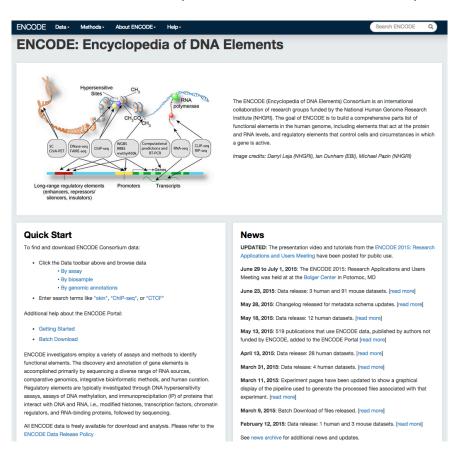


Researchers argue for standard format to cite lab resources

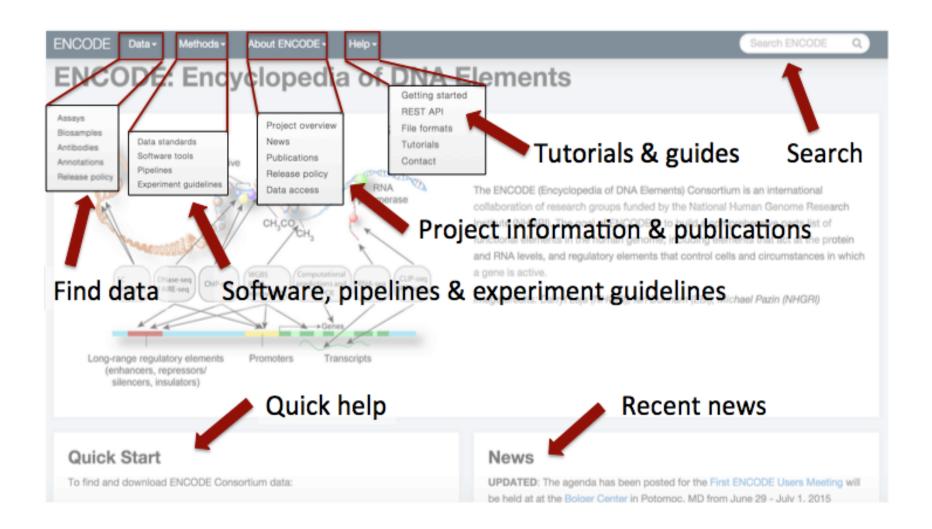
Research Resource Identifier (RRID) aims to clean up poorly referenced data. *Nature* (29 May 2015) Need for standard and unique identifiers

ENCODE portal www.encodeproject.org

- Central source for ENCODE data: experimental and analysis data
- Hub for project information: data standards & publications
- High-quality metadata: data provenance & transparency

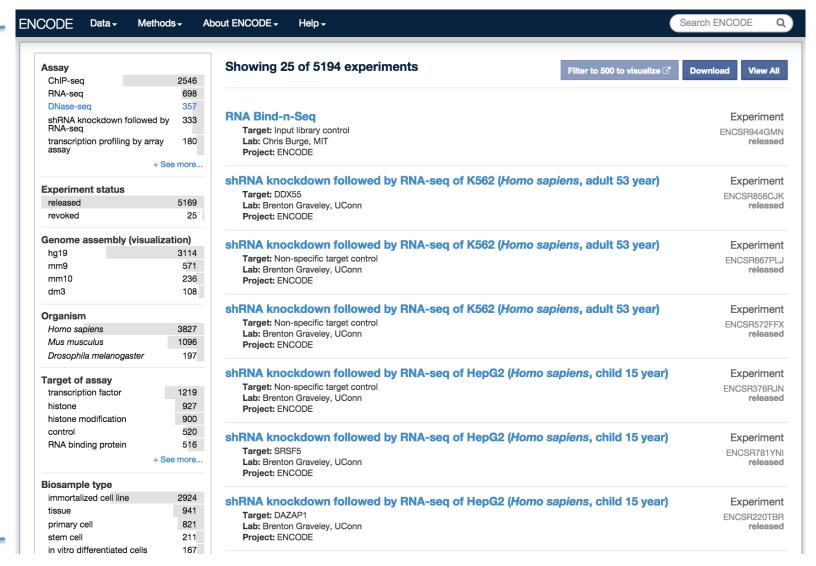


The ENCODE portal at a glance

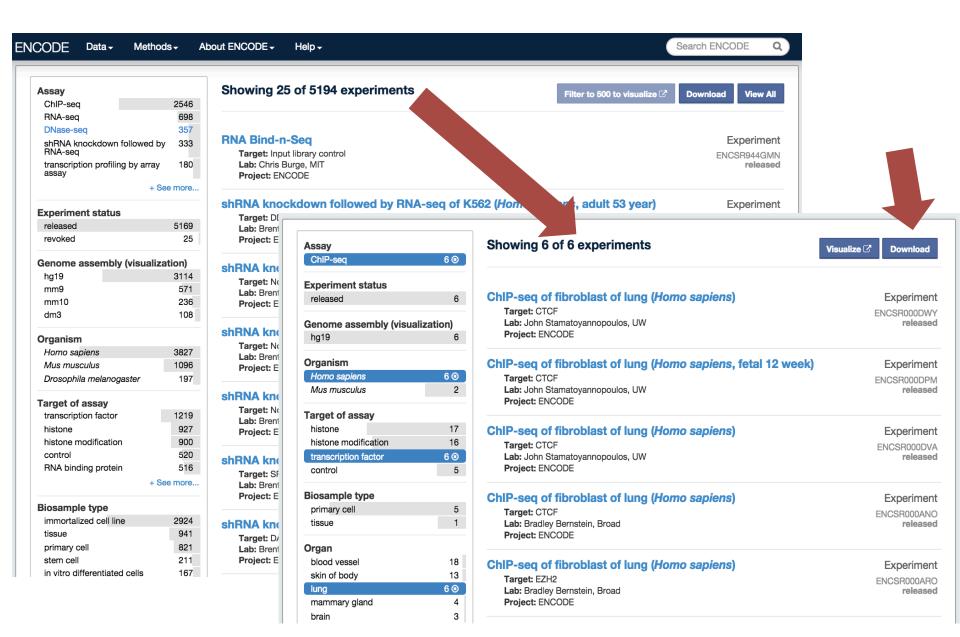


Metadata-driven faceted browsing: by assay

Select "Assay" under the "Data" menu:

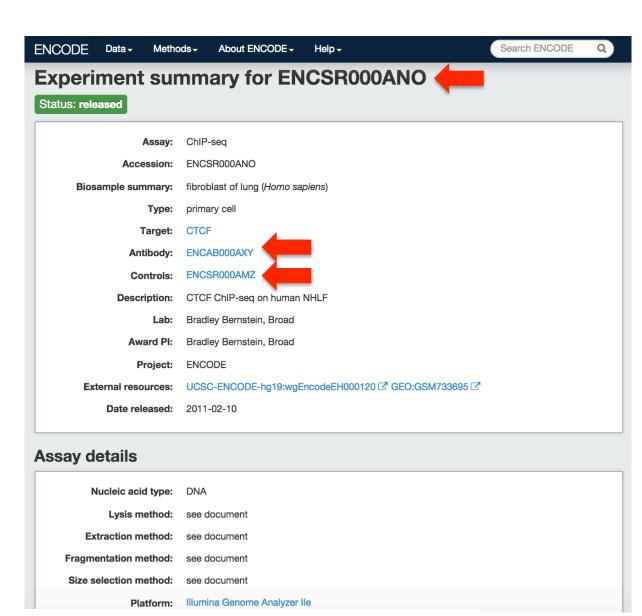


Metadata-driven faceted browsing: by assay



Main units of an experimental analysis are uniquely accessioned

Experiments – ENCSR###XXX
Biosamples – ENCBS###XXX
Donors/strains – ENCDO###XXX
Libraries – ENCLB###XXX
Antibody lots – ENCAB###XXX
Files – FNCFF###XXX



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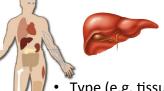
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Rich experimental metadata is collected and presented for clarity and context

For example:

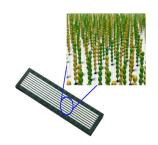
Donor & biosample

- Species
- Age
- Sex
- Health status
- Ethnicity
- Strain



- Type (e.g. tissue, cell line)
- Source
- Product id
- · Lot id
- Dates (e.g. growth, harvest,cprocurement)
- Passage number
- Starting amount
- Lab assigned IDs

Platform



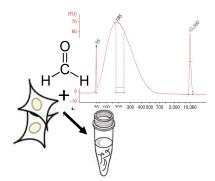
- Instrument
- Read length
- · Single or Paired end
- Lane number
- · Sequencing depth

Treatment & genetic modifications



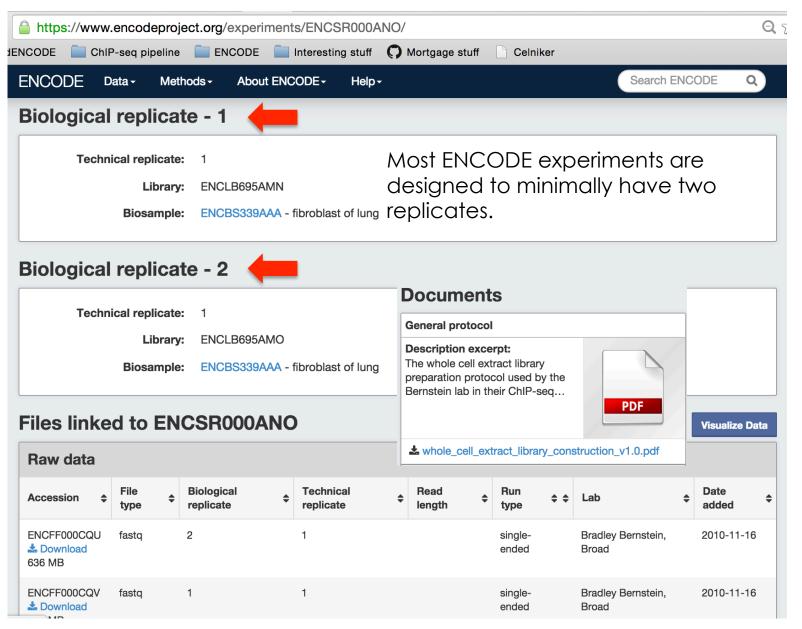
- · Agent (chemical, biological)
- Concentration
- Duration
- Construct type
- Tag
- Tag location
- · Insert sequence
- Target
- Transfection type
- Protocol

Library preparation



- · Lysis method
- · Sonication method
- Extraction method
- · Nucleic acid type
- · Nucleic acid size range
- Library preparation protocol
- Strand specificity
- Size selection method
- Validation document

Replication and transparency of methods



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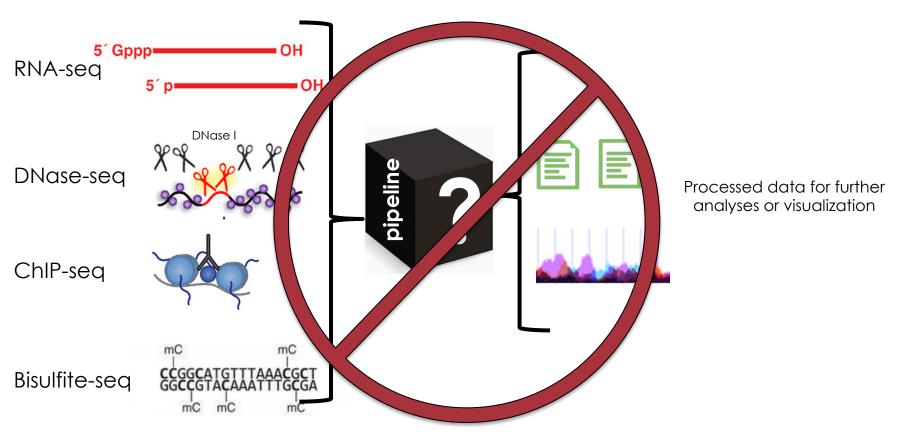


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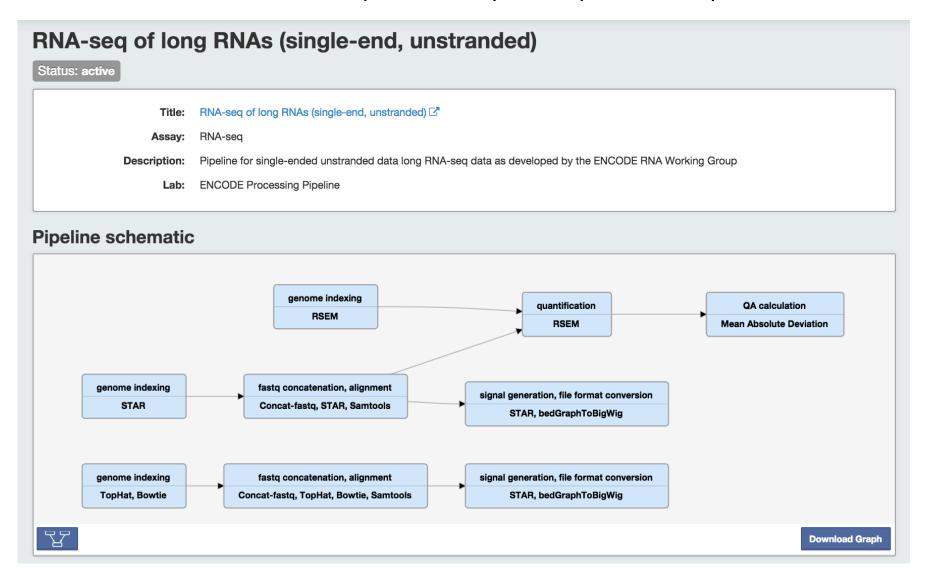


Data provenance & process transparency

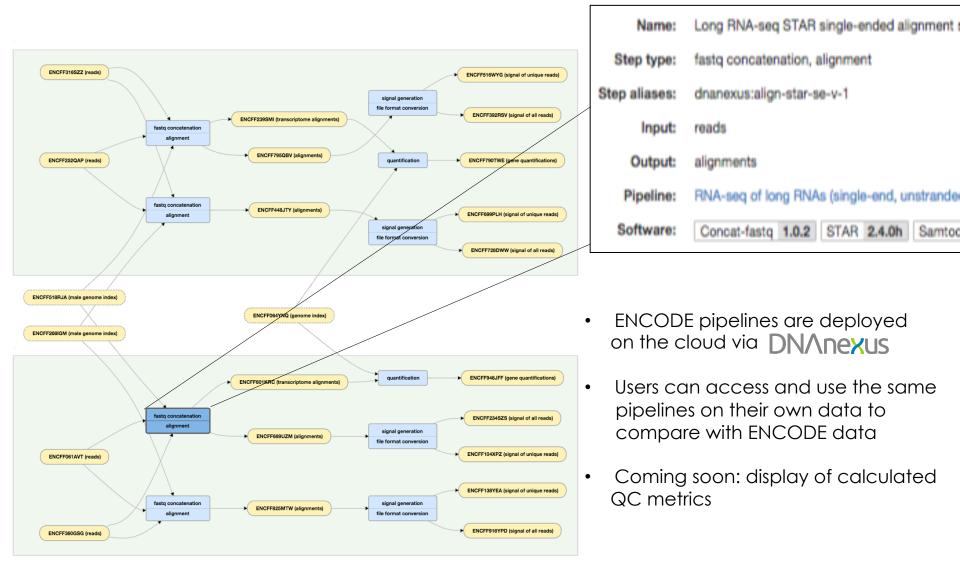


Avoid the pipeline blackbox

ENCODE runs uniform processing pipelines to enhance data comparability & replicability



Each pipeline run is tracked and viewable on the portal



Single-end long RNA-seq pipeline run for experiment ENCSR823VEE

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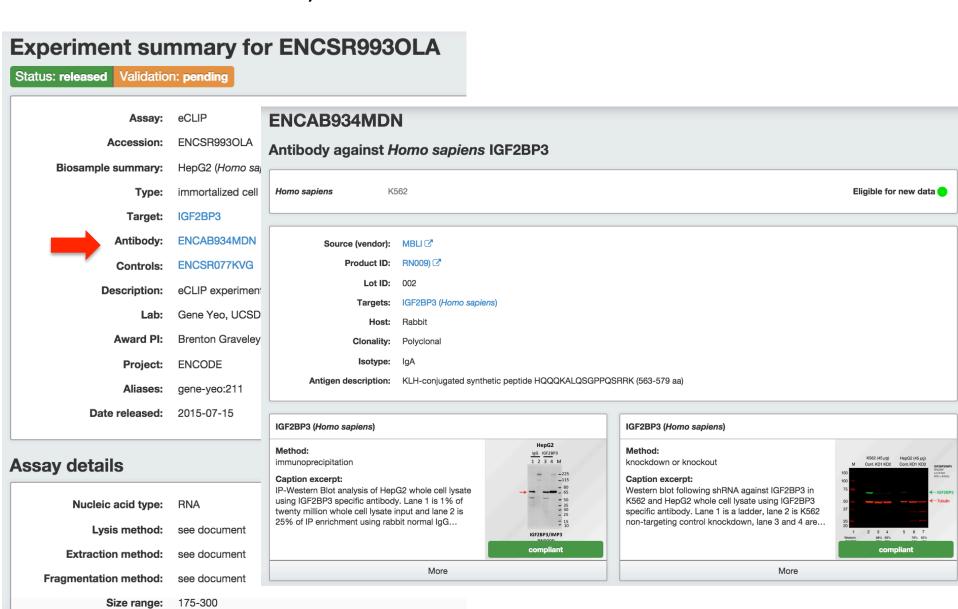


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Each antibody lot is characterized & accessioned



agarose gel extraction

Size selection method:

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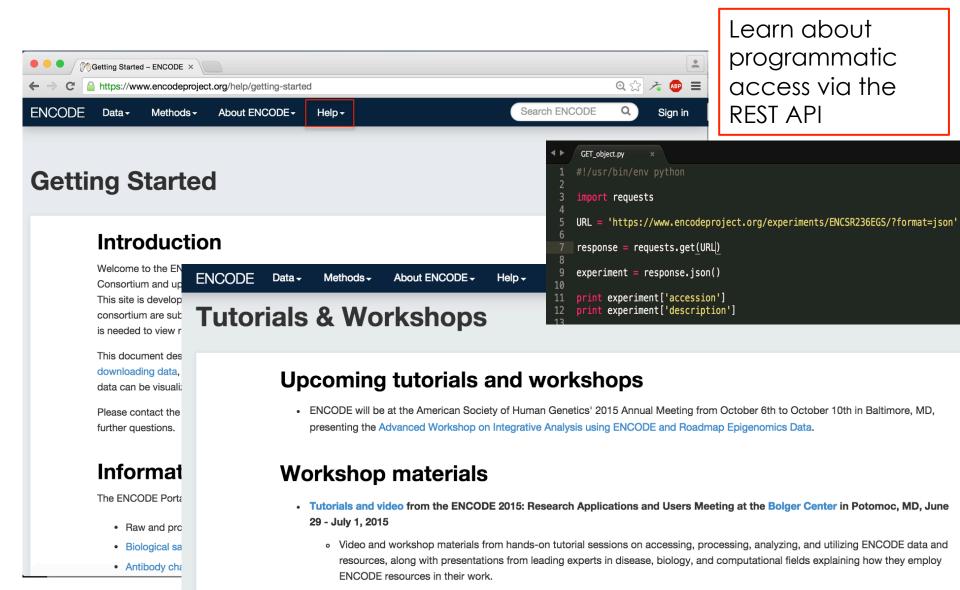


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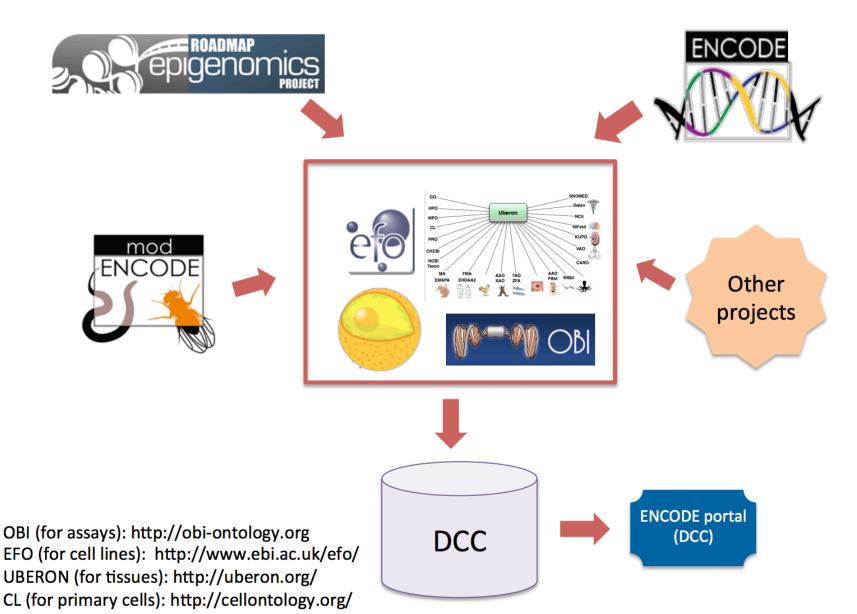
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For more details, there's help pages, tutorials, exercises and more



Metadata integration using ontologies



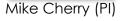
The ENCODE DCC











Ben Hitz

Cricket Sloan













Esther Chan

Jean Davidson Idan Gabdank

Seth Strattan

Marcus Ho

Aditi Narayanan

















Tim Dreszer

Marissa Melen Nikhil Podduturi Laurence Rowe Forrest Tanaka Stuart Miyasato Matt Simison

Zhenhua Wang





