



FACEBASE WORKSHOP: FAIR DATA FOR CRANIOFACIAL RESEARCH

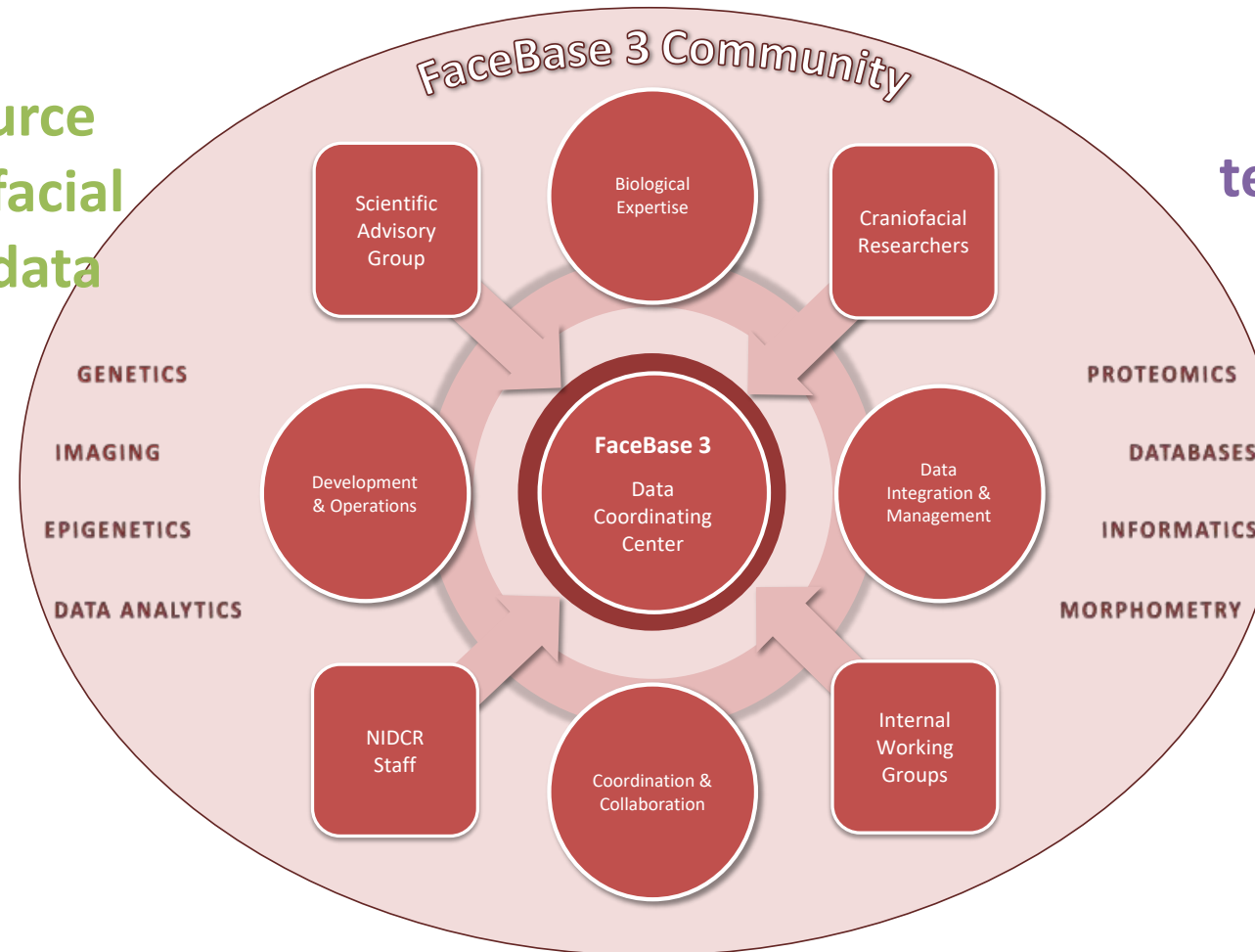
Robert Schuler

43rd Annual SCGDB Meeting

October 20, 2020

An Open Platform for Craniofacial and Dental Research

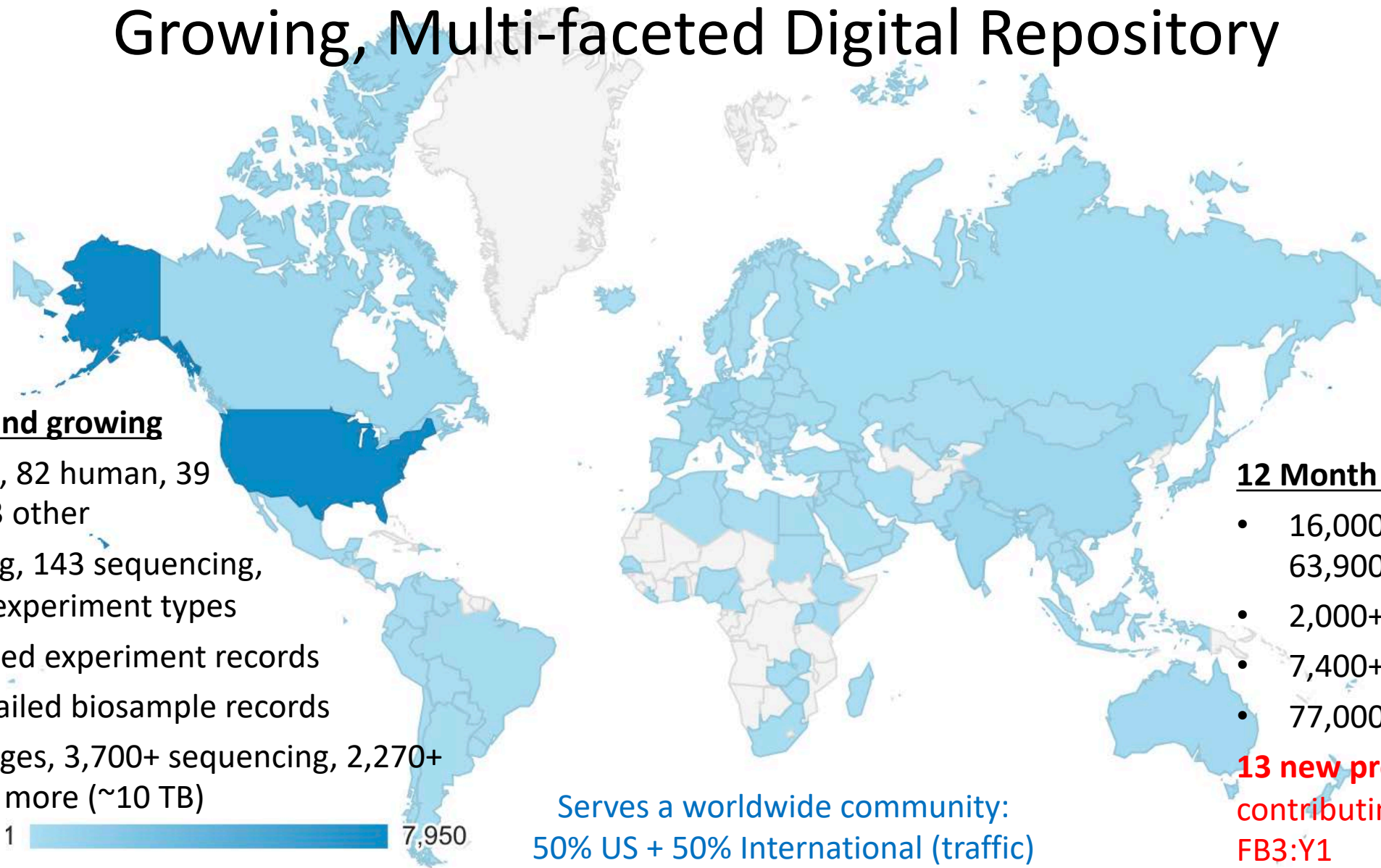
Comprehensive resource on dental and craniofacial research -- fostering data sharing and curated resources.



Comprised of a core team of craniofacial and computer science expertise w/ advisory and experts groups for oversight.

Transitioned from a “hub and spoke” consortium to a community-wide effort in 2019.

Growing, Multi-faceted Digital Repository



[illegible]

- 80% of time spent on accessing, cleaning, integrating data
- Scarcity of data sharing
- 10% reproducibility of data
- Recent high-profile retractions in COVID-19 research, for example.

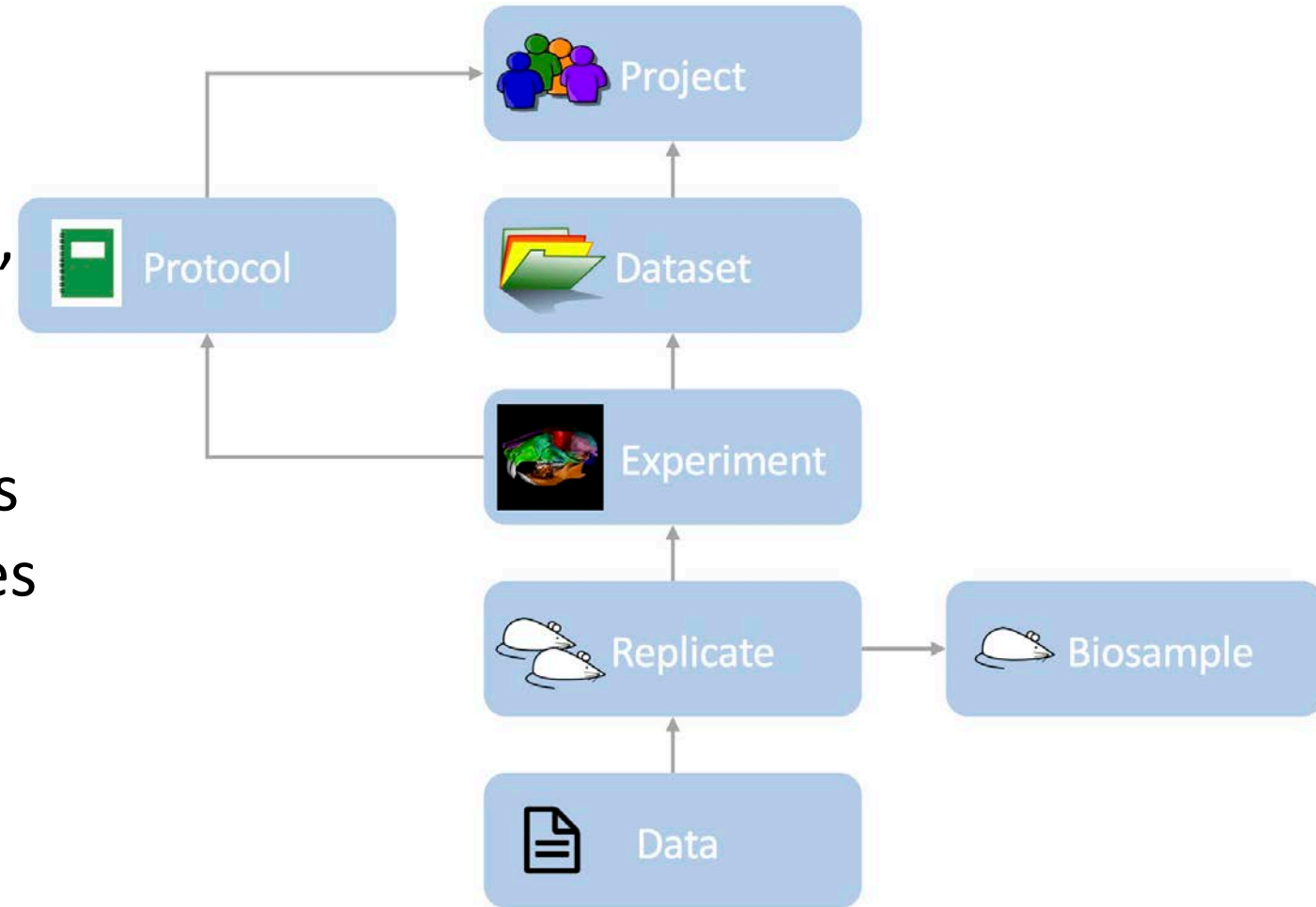
Findable **Accessible** *Interoperable* **Reusable** Data

- **Organization** of data files to understand what and how they were produced
- **Terminology** from community standards with broad acceptance
- **Description** of the experimental methods and biological materials
- **Protocols** that are enumerated or referenced for precise details



Organization of Data on FaceBase

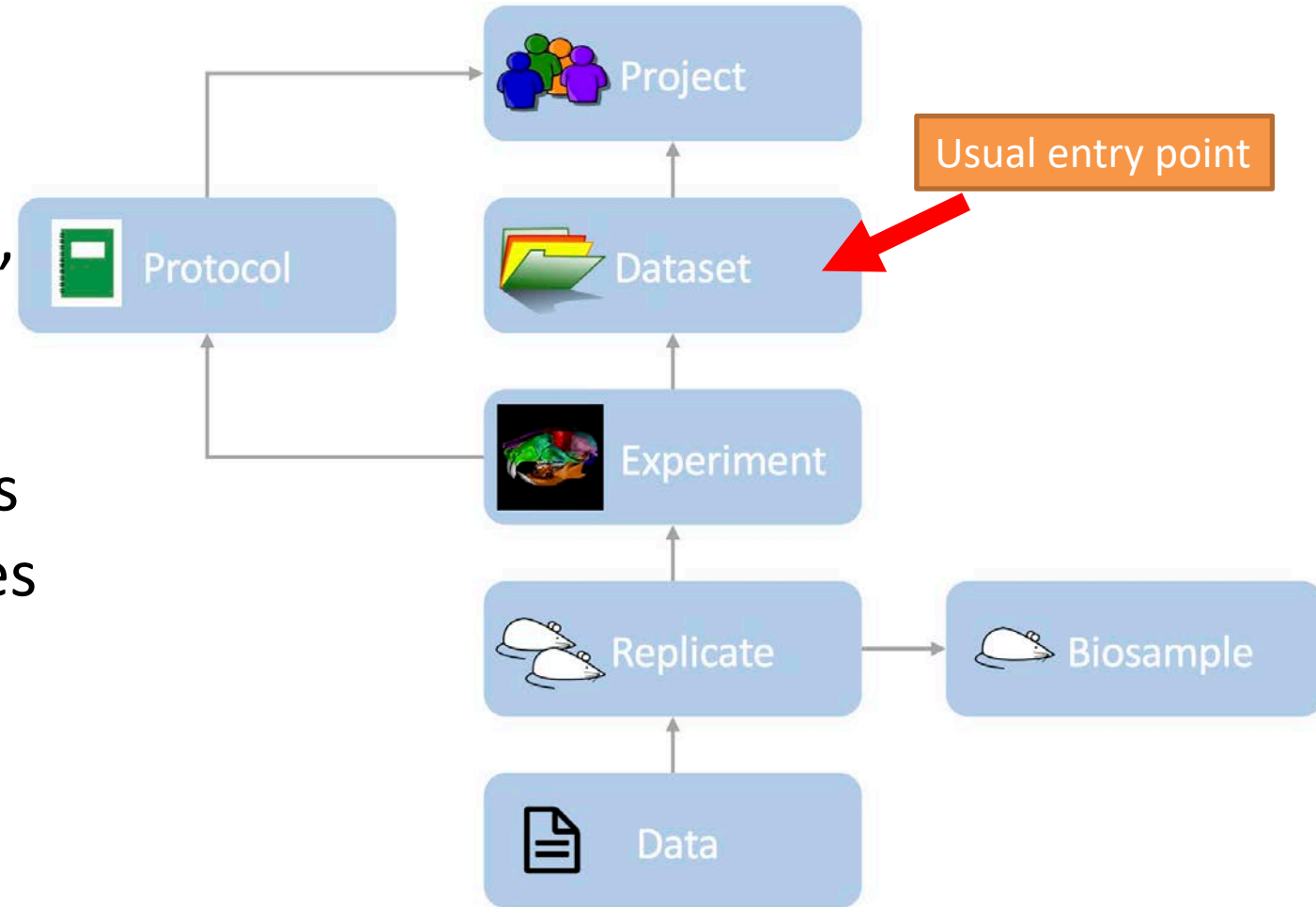
- **Projects** contribute **Datasets**
- **Datasets** organize **Experiments**, **Biosamples**, and **Data**
- **Protocols** give additional details on the materials and procedures conducted by Experiments



FaceBase data model (high-level view)

Organization of Data on FaceBase

- **Projects** contribute **Datasets**
- **Datasets** organize **Experiments**, **Biosamples**, and **Data**
- **Protocols** give additional details on the materials and procedures conducted by Experiments
- *Search generally begins at the Dataset*



FaceBase data model (high-level view)

Search and Refine

- Refinements by experiment type, species, gene, age, anatomy, etc.

The screenshot displays the FaceBase Data Browser interface. The top navigation bar includes links for Data Browser, Submit, Data Access, Resources, About, News, and Help, along with Sign Up and Log In buttons. The main section is titled "Dataset" and features a search bar and filters. The "Refine search" panel on the left shows the "Experiment Type" filter set to "rna" and the "Species" filter set to "Mus musculus". The main table displays 25 of 509 matching results, with columns for View, Record ID, Title, Experiment Type(s), Species, Stage(s), PI(s), and Release Date.

View	Record ID	Title	Experiment Type(s)	Species	Stage(s)	PI(s)	Release Date
	1-DY3G	Tooth root development in Osr2-Cre;Ezh2fl/fl and control mice	fluorescence microscopy, micro-computed tomography (microCT), microscopy assay	Mus musculus	P0, PN 2 weeks, PN 4 weeks	Yang Chai	2020-09-21
	1-71HY	Sagittal Suture, WT and Fgfr2+/S252W (Apert syndrome mouse model), E16.5 and E18.5	RNA-seq assay	Mus musculus	E16.5, E18.5, TS24, TS26	Ethilyn Wang, Jabs, Greg Peter Holmes, Harm Van Bakel	2020-08-13
	1-T8JG	Premaxillary-Maxillary Suture, WT and Fgfr2+/S252W (Apert syndrome mouse model), E16.5 and E18.5	RNA-seq assay	Mus musculus	E16.5, E18.5, TS24, TS26	Ethilyn Wang, Jabs, Greg Peter Holmes, Harm Van Bakel	2020-08-13
	1-T8KE	Squamoparietal Suture, WT and Fgfr2+/S252W (Apert syndrome mouse model), E16.5 and E18.5	RNA-seq assay	Mus musculus	E16.5, E18.5, TS24, TS26	Ethilyn Wang, Jabs, Greg Peter Holmes, Harm Van Bakel	2020-08-13
	1-TQ9T	Normalized expression data from the anterior lateral palate in E14.5 mice	RNA-seq assay	Mus musculus	E14.5	Somnath Datta	2020-07-31
	1-TQAE	Normalized expression data from the anterior medial palate in E14.5 mice	RNA-seq assay	Mus musculus	E14.5	Somnath Datta	2020-07-31
	1-TQB0	Normalized expression data from the anterior nasal palate in E14.5 mice	RNA-seq assay	Mus musculus	E14.5	Somnath Datta	2020-07-31
	1-TQBJ	Normalized expression data from the anterior oral palate in E14.5 mice	RNA-seq assay	Mus musculus	E14.5	Somnath Datta	2020-07-31
	1-TQC4	Normalized expression data from the posterior lateral palate in E14.5 mice	RNA-seq assay	Mus musculus	E14.5	Somnath Datta	2020-07-31
	1-TQCP	Normalized expression data from the posterior medial palate in E14.5 mice	RNA-seq assay	Mus musculus	E14.5	Somnath Datta	2020-07-31
	1-TQD8	Normalized expression data from the posterior lateral palate in E14.5 mice	RNA-seq assay	Mus musculus	E14.5	Somnath Datta	2020-07-31

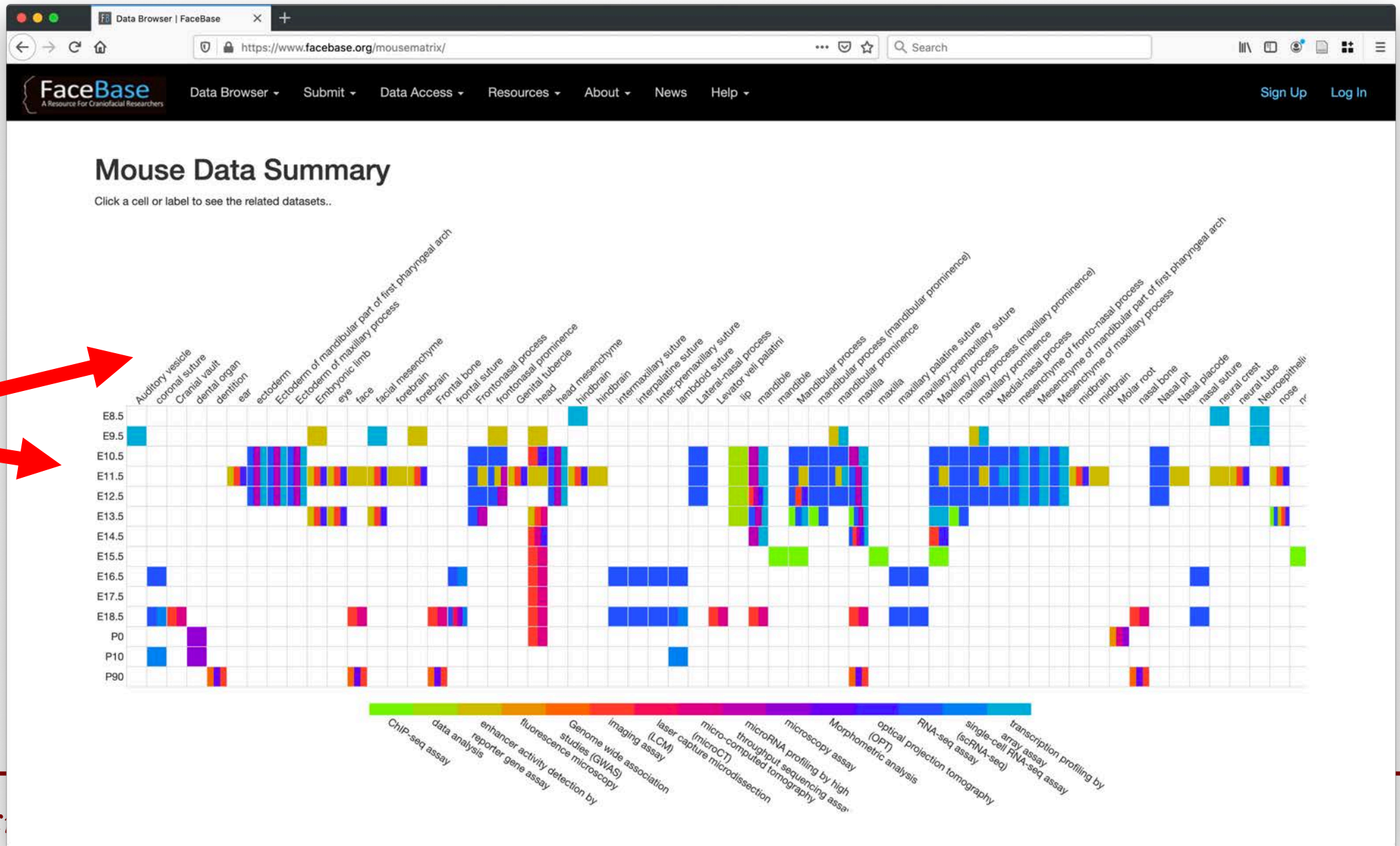
Search and Refine

- Refinements by experiment type, species, gene, age, anatomy, etc.
- Summary view with key properties listed.

The screenshot displays the FaceBase Data Browser interface. The top navigation bar includes links for Data Browser, Submit, Data Access, Resources, About, News, and Help, along with Sign Up and Log In buttons. The main section is titled 'Dataset' and features a search bar and filters for 'Experiment Type: micro-computed tomography (microCT)' and 'Species: Mus musculus'. The 'Refine search' panel on the left shows a list of experiment types with 'micro-computed tomography (microCT)' and 'RNA-seq assay' selected. The 'Species' section shows 'Mus musculus' selected. The main table displays 25 of 509 matching results, with columns for View, Record ID, Title, Experiment Type(s), Species, Stage(s), PI(s), and Release Date. Red arrows point from the text 'Refinements by experiment type, species, gene, age, anatomy, etc.' to the 'Refine search' panel and from 'Summary view with key properties listed.' to the table headers.

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Summary of Mouse Datasets



Explore By Anatomical Region of Interest

Linked to
standard
terminology

Links above
drive search
results of
related
datasets

Surface
mesh viewer
with each
structure
coded to the
terminology

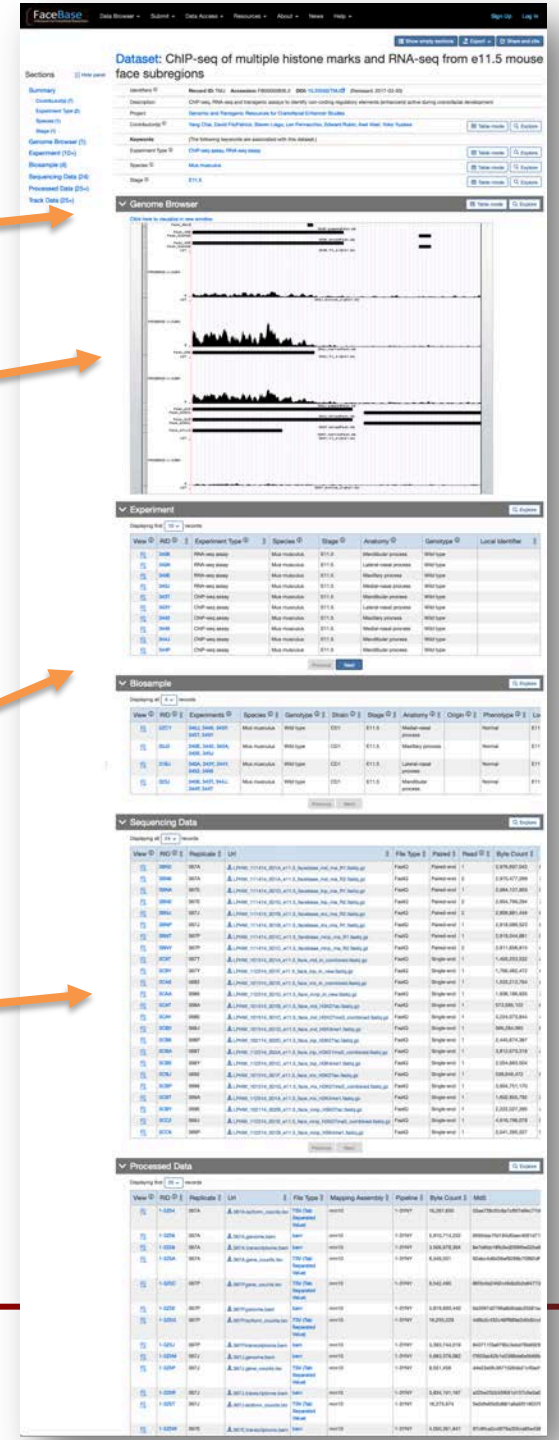
“Explore” to go
deeper into the
search and
refine actions

The screenshot shows the FaceBase web application interface. At the top, there's a browser window with the URL <https://www.facebase.org/image-nav/>. Below the browser, there's a toolbar with various interactive tools like 'Show landmarks', 'Calculate Distances', 'Show bounding box', 'Clip plane', 'Zoom in', 'Zoom out', 'Rotate', 'Reset', and 'Open'. The main content area is divided into two parts. On the left, there's a 'Meshes' panel with a list of anatomical structures: frontal suture, interparietal bone, mandible, maxilla, nasal bone, occipital bone, palatine bone, Parietal bone, premaxilla, and skull. Each item has a small icon and a color-coded box. On the right, there's a 3D surface mesh viewer showing a mouse skull with different parts colored to match the mesh list. Below the mesh viewer, there's a section titled 'Anatomy[®]: maxilla' with a 'Summary' link and a 'Datasets (21)' link. The 'Datasets' section is expanded, showing a table of 21 records. The table has columns for View, Record ID, Title, Experiment Type(s), Species, Stage(s), and Contributor. The first three records are visible.

View	Record ID	Title	Experiment Type(s)	Species	Stage(s)	Contributor
	1-731C	Facial shape and allometry quantitative trait loci in the Diversity Outbred mouse	Genome wide association studies (GWAS), imaging assay, Morphometric analysis	Mus musculus	P90	Benedikt H. Charles C. F. David Katz, Cheverud, J. Fish, Ralph Marcucio, F. Green, Trev
	1-517M	Interactive Zebrafish Craniofacial Atlases	micro-computed tomography (microCT)	Danio rerio	adult	Azman Rasi Katherine W. Matthew H. Shannon Fi
	1-4TDT	Whole-genome transcriptome profiling of E13.5 Wnt1-Cre:Alk5 ^{fl/fl} hv	transcription profiling by array assay	Mus musculus	E13.5	Paul Thome Sanchez, Yi

Overview of Dataset Display

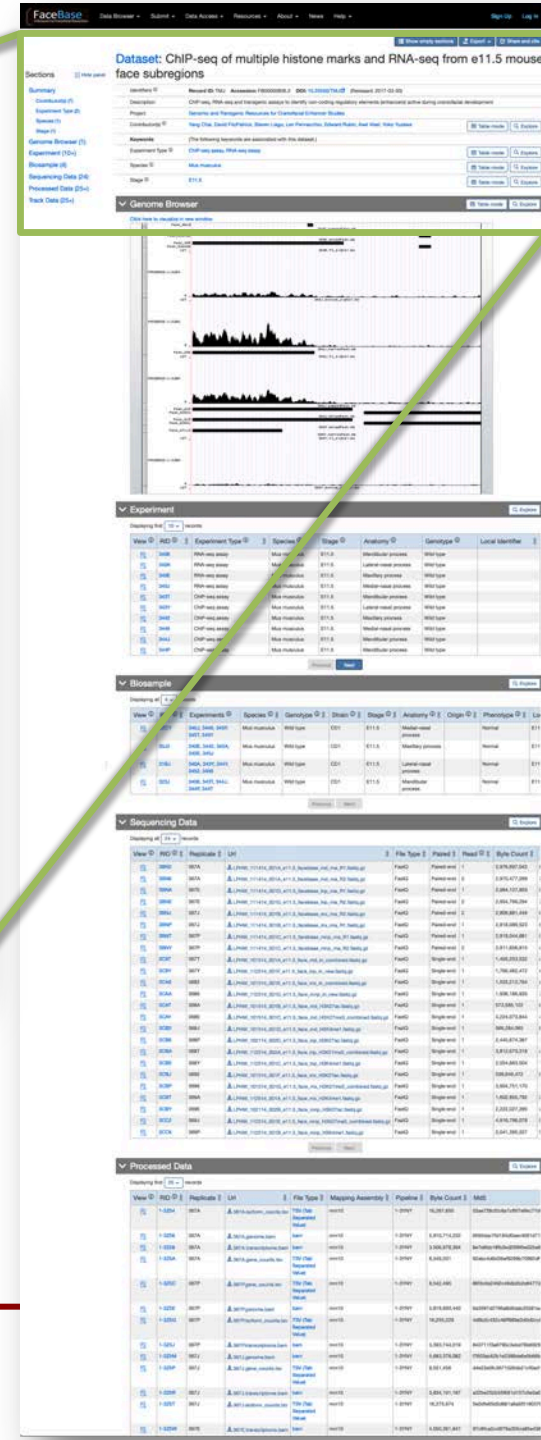
- Summary
- Visualization
 - Genome Browser
 - Surface Models
 - Orthoslice Views
 - Images
- Experiments & Biosamples
- Data Files
 - Sequencing, Processed, Imaging, Array, Supplementary, other...



Summary of the Dataset Contents

Dataset: ChIP-seq of multiple histone marks and RNA-seq from e11.5 mouse face subregions

Identifiers ^①	Record ID: TMJ Accession: FB00000806.2 DOI: 10.25550/TMJ (Released: 2017-03-30)	
Description	ChIP-seq, RNA-seq and transgenic assays to identify non-coding regulatory elements (enhancers) active during craniofacial development	
Project	Genomic and Transgenic Resources for Craniofacial Enhancer Studies	
Contributor(s) ^①	Yang Chai, David FitzPatrick, Steven Lisgo, Len Pennacchio, Edward Rubin, Axel Visel, Yoko Yuzawa	Table mode Explore
Keywords	(The following keywords are associated with this dataset.)	
Experiment Type ^①	ChIP-seq assay, RNA-seq assay	Table mode Explore
Species ^①	Mus musculus	Table mode Explore
Stage ^①	E11.5	Table mode Explore

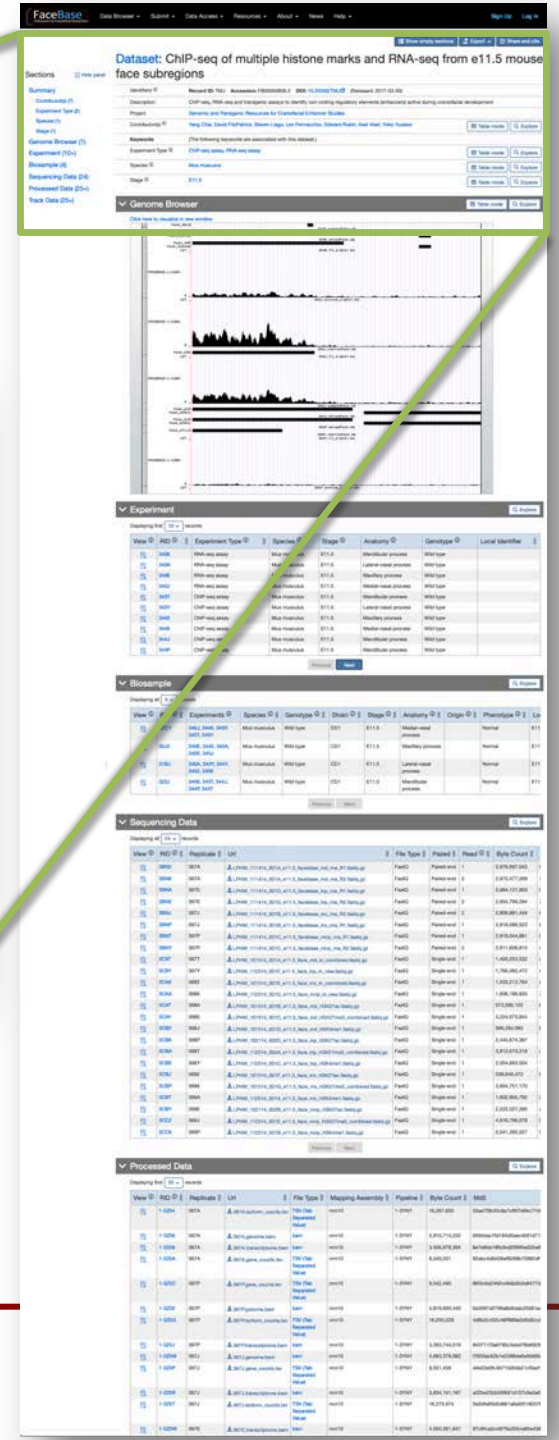


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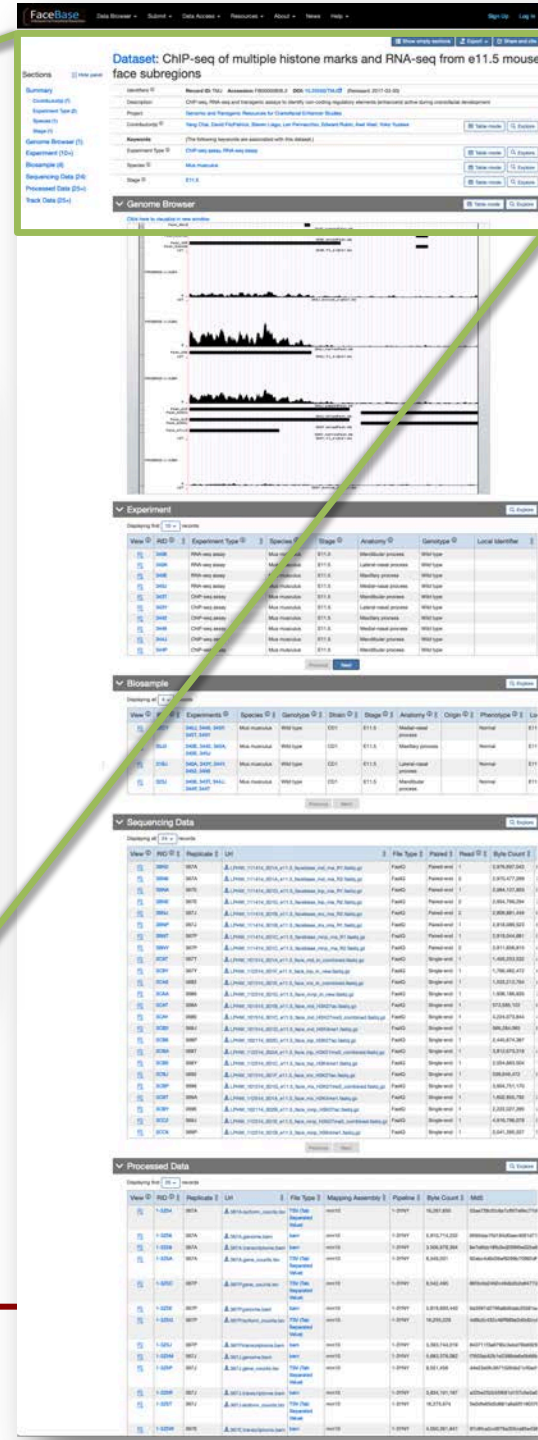
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Attribution of contributors



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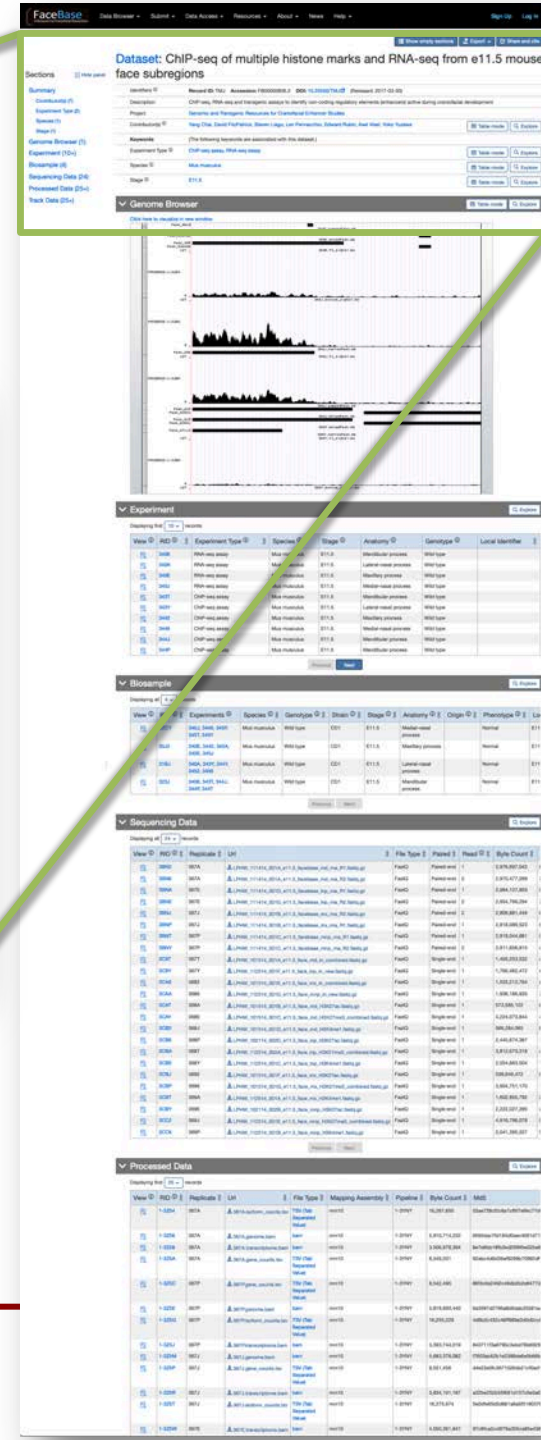
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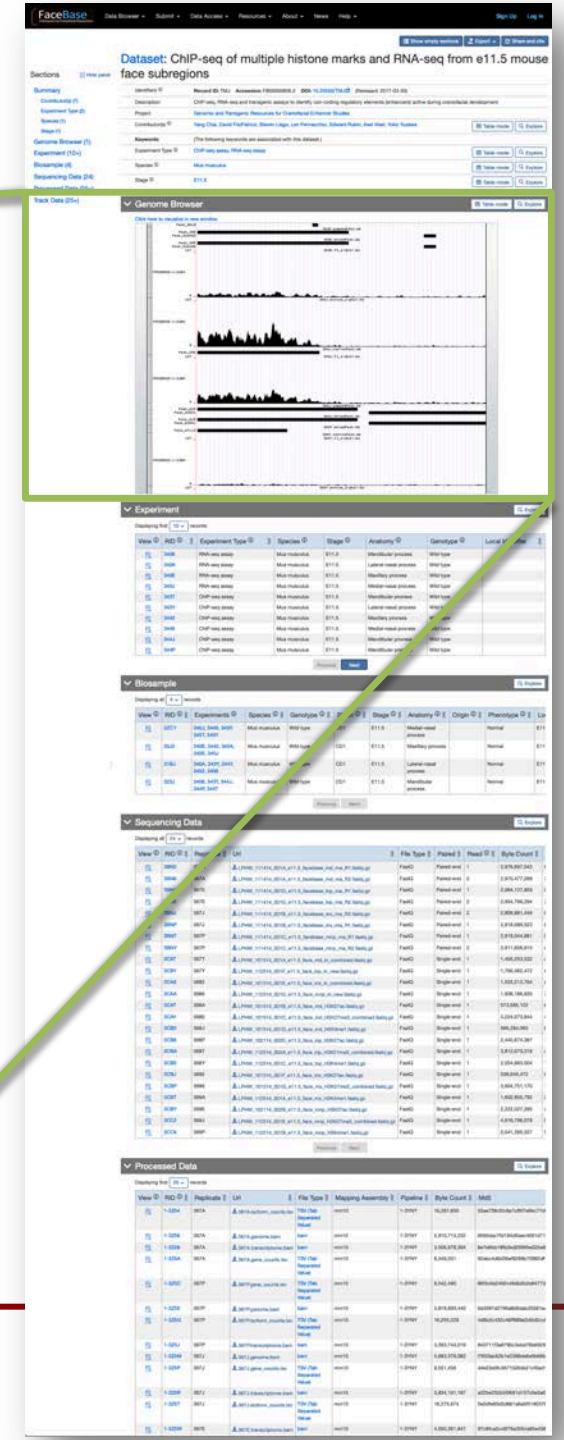
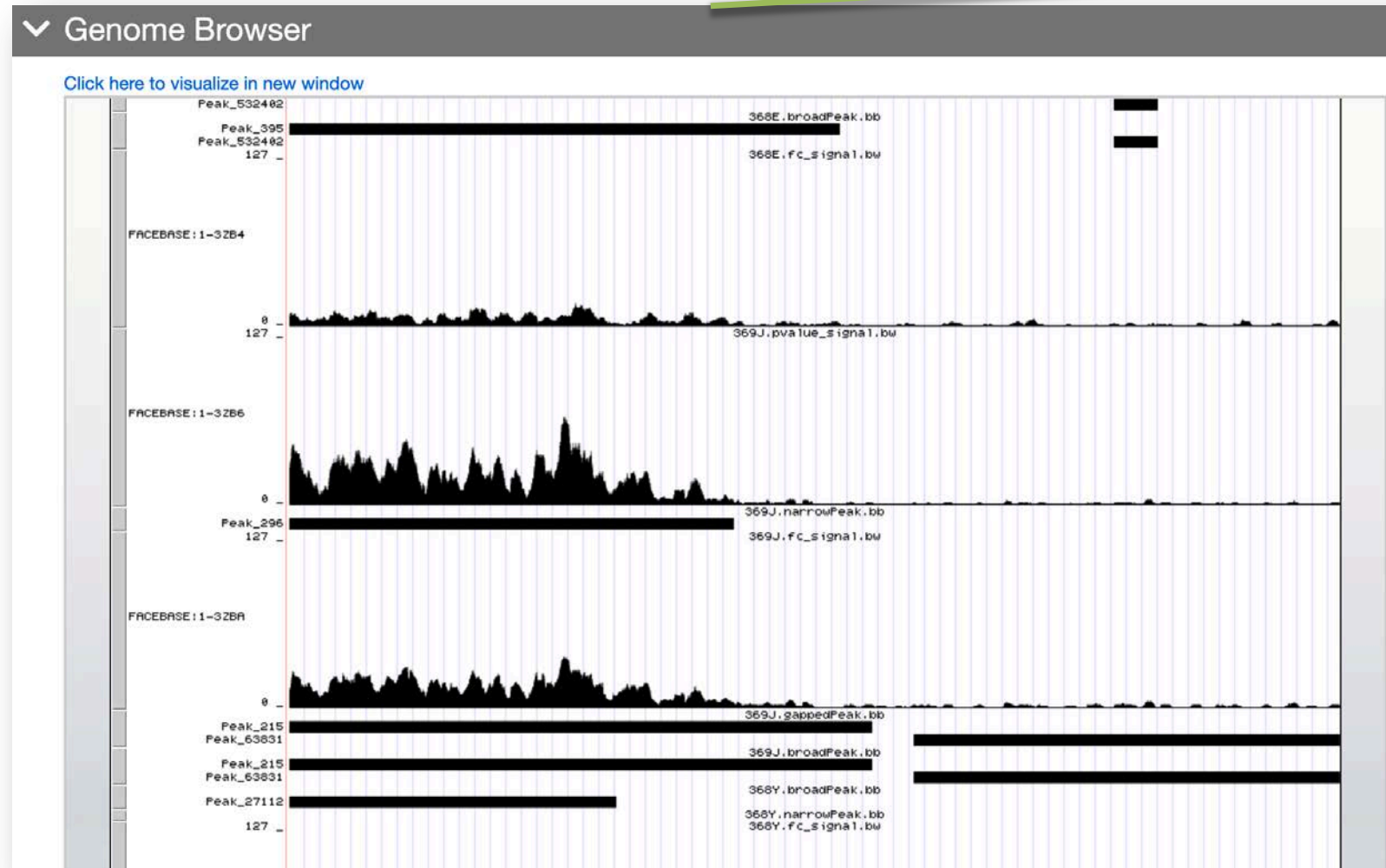
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Stage ^①	E11.5

Keywords from standardized terminology

Attribution of contributors



Embedded Visualization



Integrated Genome Browser Track Hub

FaceBase
disseminated
on Genome
Browser

FaceBase
Data Browser Submit Data Access Resources About News Help Sign Up Log In

Dataset: ChIP-seq of multiple histone marks and RNA-seq from e11.5 mouse face subregions

Sections Hide panel

- Summary
- Contributors (7)
- Species (1)
- Experiment Type (2)
- Stage (1)
- Genome Browser (1)
- Experiment (20)
- Biosample (4)
- Sequencing Data (24)
- Processed Data (25+)
- Track Data (25+)

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Project	Vitel: Genomic and Transgenic Resources for Craniofacial Enhancer Studies
Species	Mus musculus
Experiment Type	ChIP-seq assay RNA-seq assay
Stage	E11.5

Genome Browser

Click here to visualize in new window

UCSC Genome Browser on Mouse Dec. 2011 (GRCm38/mm10) Assembly

chr12:56,694,976-56,714,605 19,630 bp

enter position, gene symbol or search terms go

Scale chr12: 127 56,695,000 5 kb 56,700,000 56,705,000

FACEBASE:1-3ZAC

FACEBASE:1-3ZB4

FACEBASE:1-3ZB6

FACEBASE:1-3ZB8

FACEBASE:1-3ZB9

FACEBASE:1-3ZC0

FACEBASE:1-3ZC1

FACEBASE:1-3ZC2

FACEBASE:1-3ZC3

FACEBASE:1-3ZC4

FACEBASE:1-3ZC5

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FACEBASE:1-3ZZ4

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FACEBASE:1-3ZZ9

Embedded Browser

1

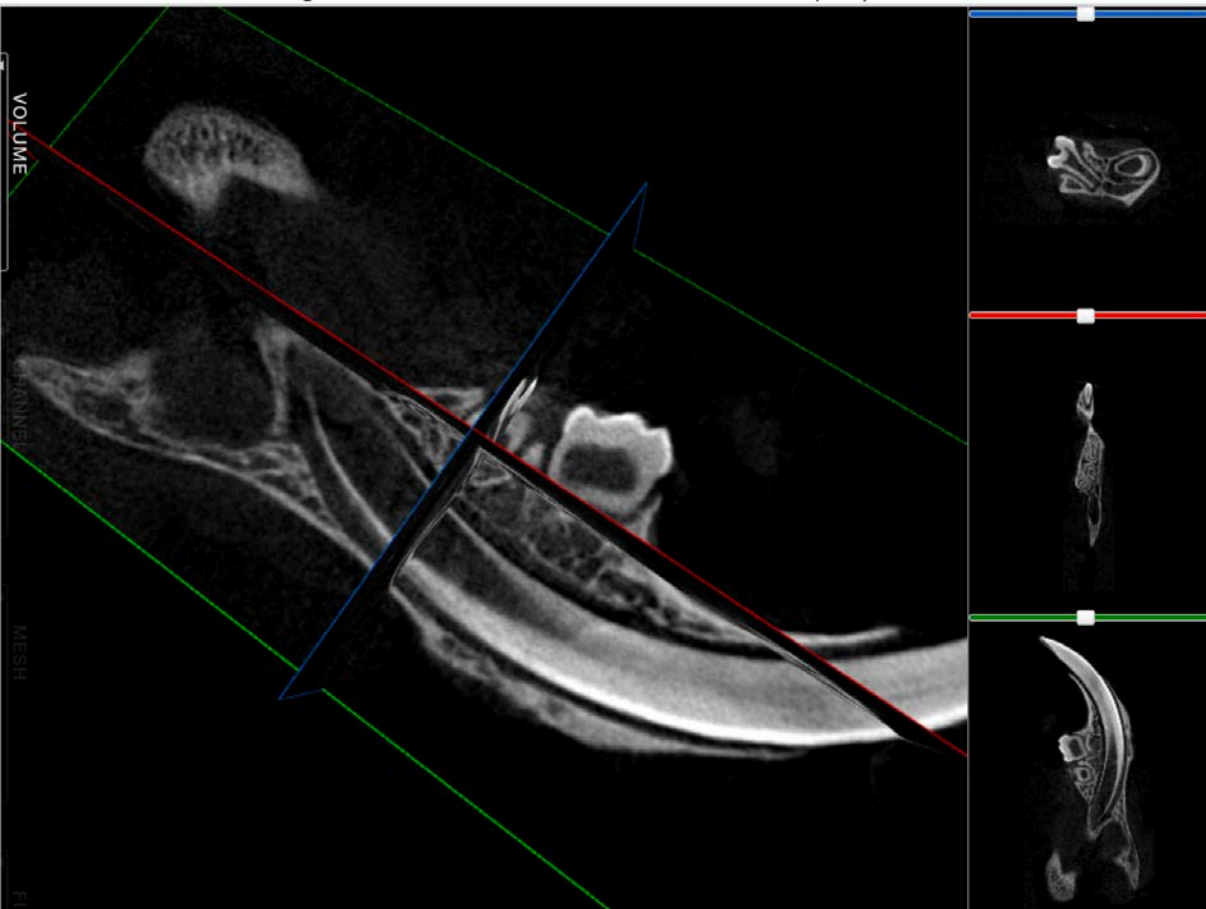
2

3

Tracks linked
back to FaceBase

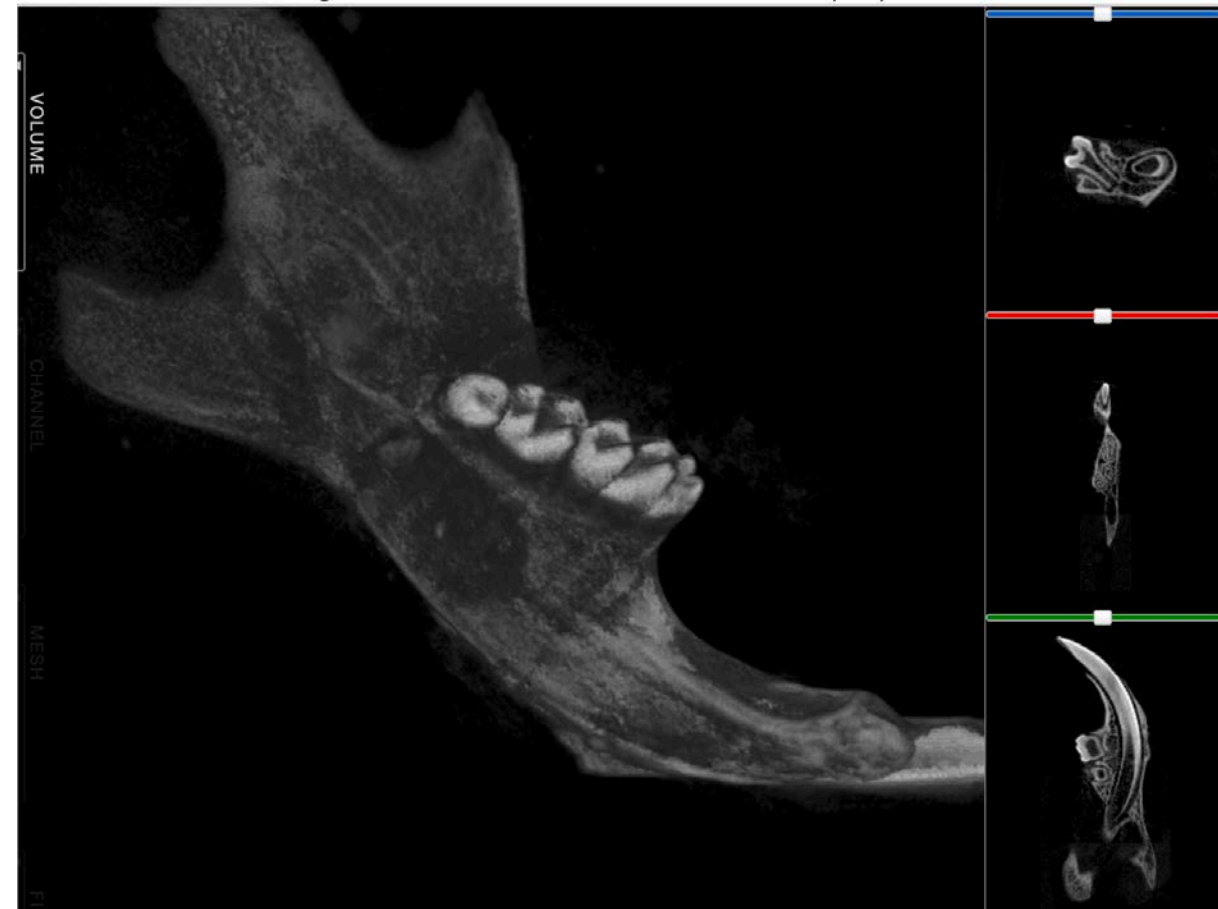
Orthogonal Slice Viewer with Volume Rendering

JJ-Osr2-Ezh2-mut-4wks-1.nii.gz – click the load button below to view the downsampled preview



Orthoslice View

JJ-Osr2-Ezh2-mut-4wks-1.nii.gz – click the load button below to view the downsampled preview

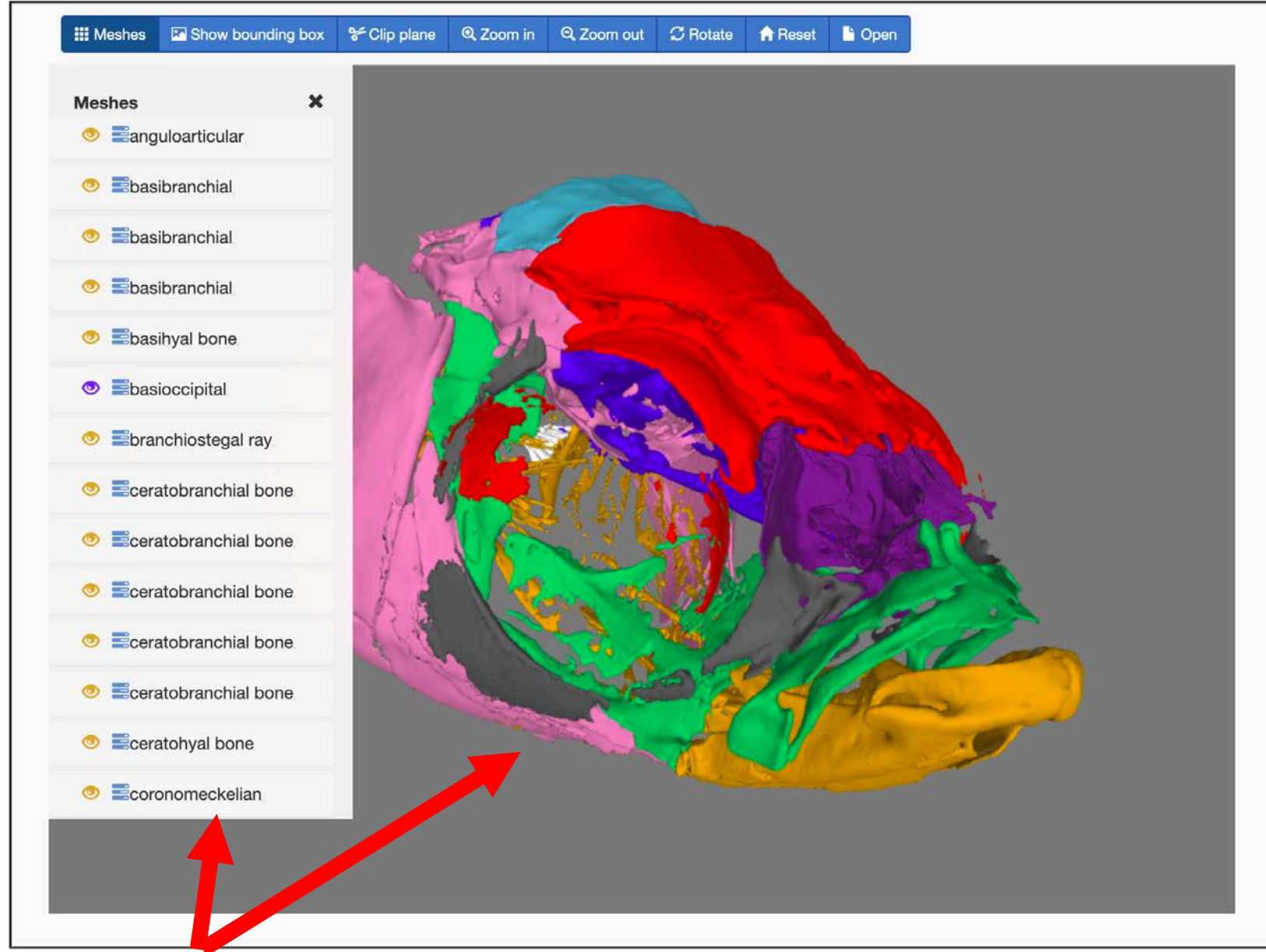


3D Volume View

Web-based Viewer

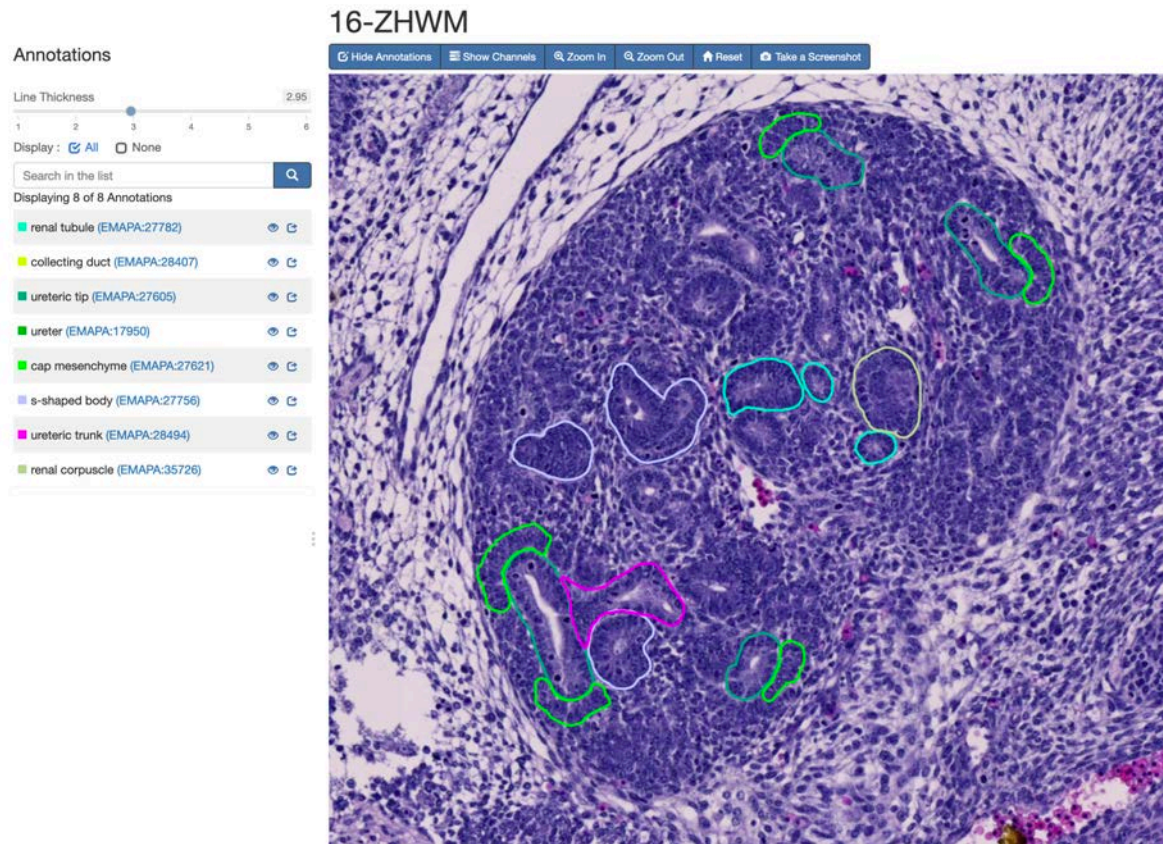
Highly Detailed Surface Models

- Surface models may contain any number of anatomical structures
- Linked to underlying (meta)data
- Linked to standard vocabulary (ZFA, MA, UBERON,...)

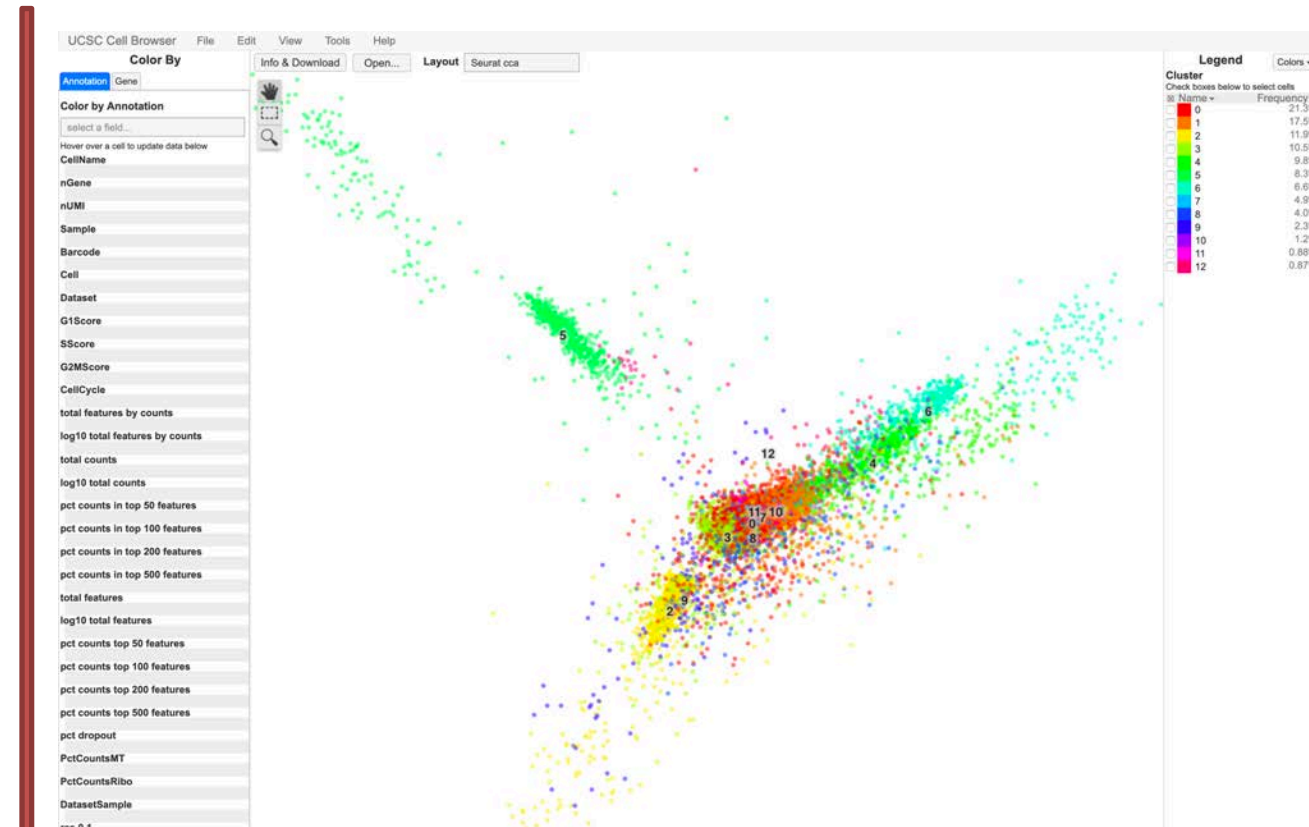


60+ anatomical structures linked to standard terminology in Zebrafish atlas (pictured)

Upcoming Visualization Features



**High-resolution Microscopy
Images with Annotations**













**Single Cell Browser through
Integration with UCSC**

Details on Experiments, Biosamples, etc.

Experiment

Explore

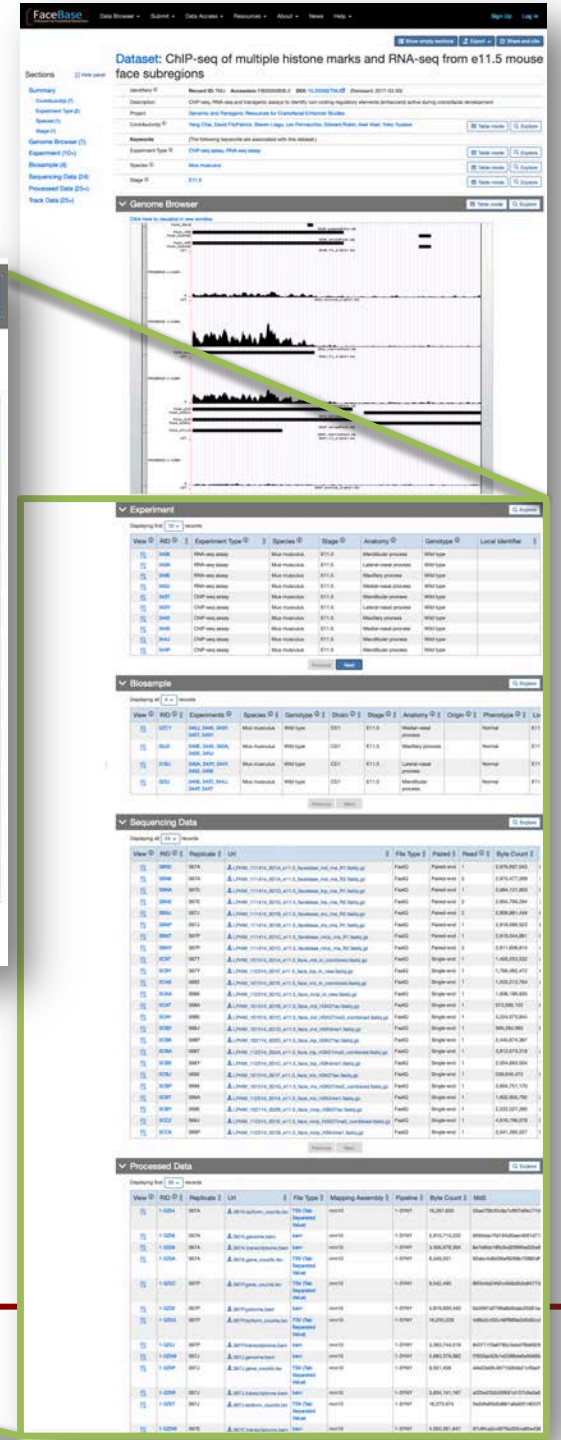
Displaying first 10 records

View	RID	Experiment Type	Species	Stage	Anatomy	Genotype	Local Identifier
	3406	RNA-seq assay	Mus musculus	E11.5	Mandibular process	Wild type	
	340A	RNA-seq assay	Mus musculus	E11.5	Lateral-nasal process	Wild type	
	340E	RNA-seq assay	Mus musculus	E11.5	Maxillary process	Wild type	
	340J	RNA-seq assay	Mus musculus	E11.5	Medial-nasal process	Wild type	
	343T	ChIP-seq assay	Mus musculus	E11.5	Mandibular process	Wild type	
	343Y	ChIP-seq assay	Mus musculus	E11.5	Lateral-nasal process	Wild type	
	3442	ChIP-seq assay	Mus musculus	E11.5	Maxillary process	Wild type	
	3446	ChIP-seq assay	Mus musculus	E11.5	Medial-nasal process	Wild type	
	344J	ChIP-seq assay	Mus musculus	E11.5	Mandibular process	Wild type	
	344P	ChIP-seq assay	Mus musculus	E11.5	Mandibular process	Wild type	

Previous

Next

Further drill-down to related biosamples, files, and other related data.



Export (i.e., Download) Dataset

Dataset: ChIP-seq of multiple histone marks and RNA-seq from e11.5 mouse face subregions

Identifiers ^①	Record ID: TMJ Accession: FB00000806.2 DOI: 10.25550/TMJ (Released: 2017-03-30)
Description	ChIP-seq, RNA-seq and transgenic assays to identify non-coding regulatory elements (enhancers) active during craniofacial development
Project	Genomic and Transgenic Resources for Craniofacial Enhancer Studies
Contributor(s) ^①	Yang Chai, David FitzPatrick, Steven Ligo, Len Pennacchio, Edward Rubin, Axel Visel, Yoko Yuzawa
Keywords	(The following keywords are associated with this dataset.)
Experiment Type ^①	ChIP-seq assay, RNA-seq assay
Species ^①	Mus musculus
Stage ^①	E11.5

Show empty sections Export Share and cite

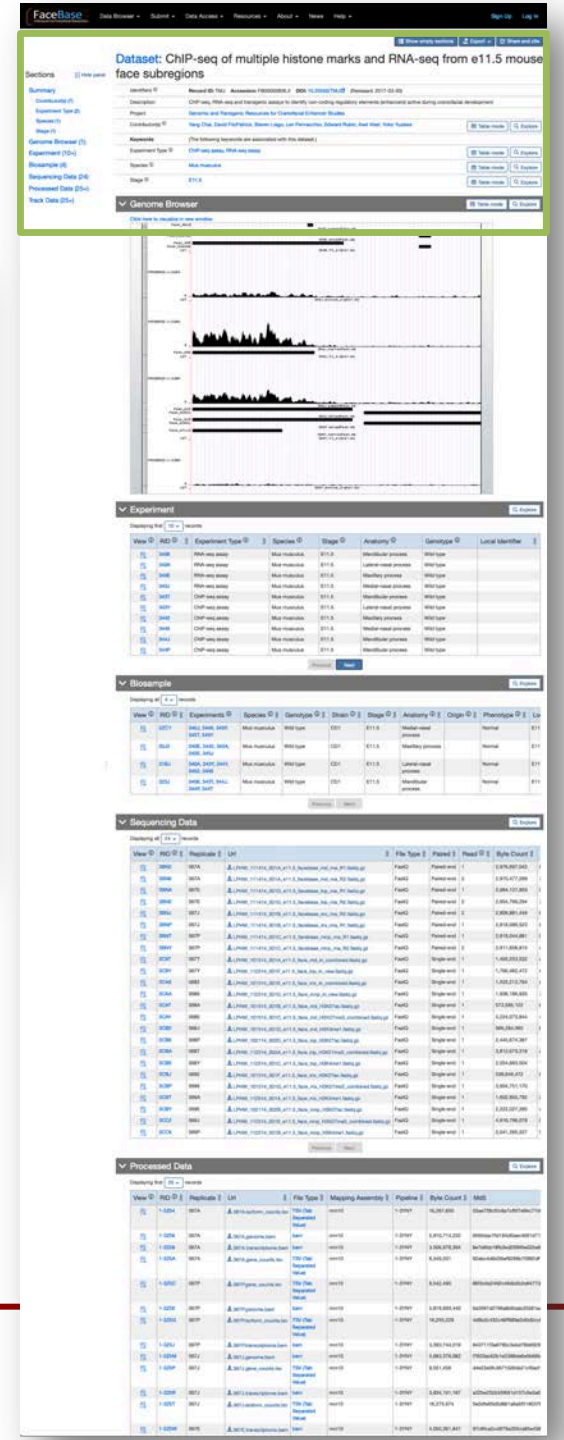
search results (csv)
BAG

Exporting BAG

Your request is being processed...

You will be prompted to download the file when it is ready.

Cancel



“Big Data Bag” Format for Large Datasets

Dataset: ChIP-seq of multiple histone marks and RNA-seq from e11.5 mouse face subregions

Identifiers ①

Description

Project

Contributor(s) ①

Keywords

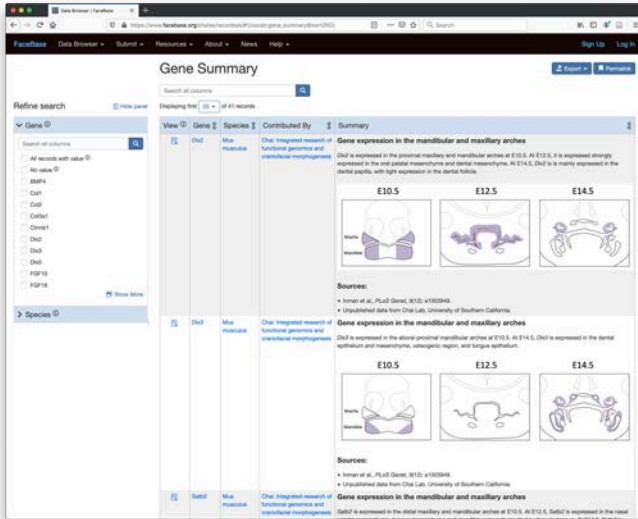
Experiment Type ①

Species ①

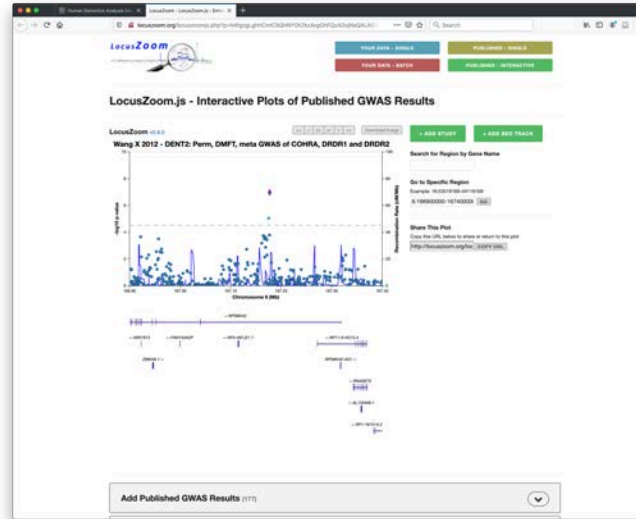
Stage ①

Name	Date Modified	Size	Kind
bag-info.txt	Today at 3:24 PM	303 bytes	Plain Text
bagit.txt	Today at 3:24 PM	55 bytes	Plain Text
data	Today at 3:24 PM	--	Folder
Biosample.csv	Today at 3:24 PM	1 KB	CSV Document
Contributor(s).csv	Today at 3:24 PM	433 bytes	CSV Document
Dataset.csv	Today at 3:24 PM	262 bytes	CSV Document
Experiment Type.csv	Today at 3:24 PM	559 bytes	CSV Document
Experiment.csv	Today at 3:24 PM	5 KB	CSV Document
Genome Browser.csv	Today at 3:24 PM	186 bytes	CSV Document
Processed Data.csv	Today at 3:24 PM	41 KB	CSV Document
Quality Control Issues.csv	Today at 3:24 PM	151 bytes	CSV Document
Sequencing Data.csv	Today at 3:24 PM	8 KB	CSV Document
Species.csv	Today at 3:24 PM	157 bytes	CSV Document
Stage.csv	Today at 3:24 PM	150 bytes	CSV Document
Track Data.csv	Today at 3:24 PM	30 KB	CSV Document
fetch.txt	Today at 3:24 PM	43 KB	Plain Text
manifest-md5.txt	Today at 3:24 PM	22 KB	Plain Text
metadata	Today at 3:24 PM	--	Folder
manifest.json	Today at 3:24 PM	120 KB	JSON Document
tagmanifest-md5.txt	Today at 3:24 PM	238 bytes	Plain Text

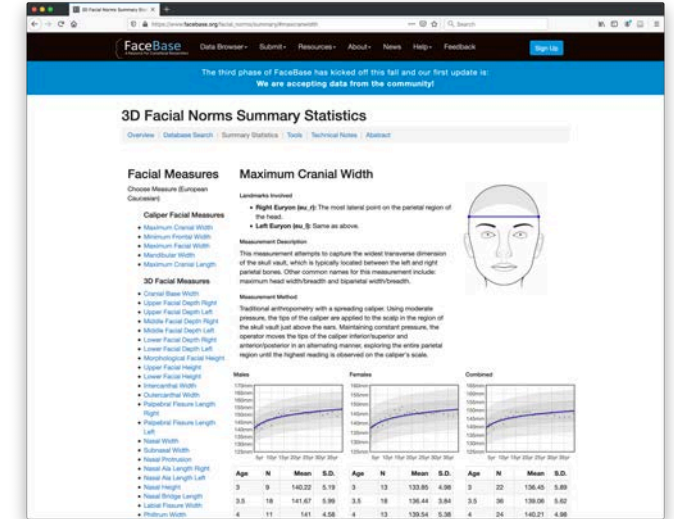
Spreadsheets (CSV)
File Manifest (links + checksums)
Client tool for reliable download



Detailed gene summaries



Interactive plots of GWAS data



Facial norms measures

More at the Resources Hub...

<https://www.facebase.org/resources/>

FaceBase Data Browser Resources About Projects News Help Sign Up Log In

Single Cell Sequencing - Coronal Suture Dataset

Warning Your login session has expired. You are not logged in.

Record ID	3TYP
Accession	FB00000970
DOI	10.25550/3TYP
Title	Single Cell Sequencing of the Coronal Suture
Description	Single Cell RNA-Seq libraries were prepared after protease digestion of the coronal suture. Libraries are available for the coronal suture. All libraries are available for the coronal suture. Single Cell Gene Expression (Version 2) K

• Greg Holmes

Share

X Close

Share Link

Versioned Link (19 hours ago)

<https://www.facebase.org/id/3TYP@2V3-Z49Y-6970>

Live Link

<https://www.facebase.org/id/3TYP>

Data Citation

Greg Peter Holmes, Ethylin Wang Jabs, Harm Van Bakel. Single Cell Sequencing - Coronal Suture, Wild Type, E18.5 and P10. *FaceBase Consortium* <https://doi.org/10.25550/3TYP> (2018).

Download Data Citation:

[BibTex](#)

Contents

- Main
- Contributors (3)
- Organism (1)
- Experiment Type (1)
- Stage (3)
- Anatomy (1)
- Mouse Genetic Background (1)**
- Genotype (1)

Why Share Your Data?

- Dedicated focus on craniofacial and dental
- Increase the visibility and impact of your research
- Cross-reference with publication
- Data are “published” like first-class academic works
- Satisfy data sharing requirements for grants and publications

Data Citation

- Persistent actionable identifiers for *all database entities*
- Explore any **historical version** of *entire database*

Follows leading publisher's recommendations

Import into reference managers

Share [X Close]

Share Link

Versioned Link (19 hours ago) **Persistent, actionable identifiers**

<https://www.facebase.org/id/2ARP@2V3-Z49Y-6970>

Live Link **Versioned or "live"**

<https://www.facebase.org/id/2ARP>

Data Citation

Greg Peter Holmes, Ethylin Wang Jabs, Harm Van Bakel. Intermaxillary suture, WT and Fgfr2+/S252W (Apert syndrome mouse model), E16.5 and E18.5. *FaceBase Consortium* <https://doi.org/10.25550/2ARP> (2018). **Digital Object Identifiers**

Download Data Citation:

[BibTex](#)

Identifiers ①	Record ID: 2A
Description	RNA-Seq library development and maxillary t
Project	Transcriptome
Contributor(s) ①	Greg Holmes,
Keywords	(The following
Experiment Type ①	RNA-seq ass
Species ①	Mus musculus
Gene ①	Fgfr2
Stage ①	E16.5, E18.5, TS24, TS26
Anatomy ①	intermaxillary suture
Mouse Genetic Background ①	C57BL/6J
Gender ①	male organism
Genotype ①	Fgfr2+/S252W; E1A-Cre, Wild type

Empower Contributors to Submit Valuable Data

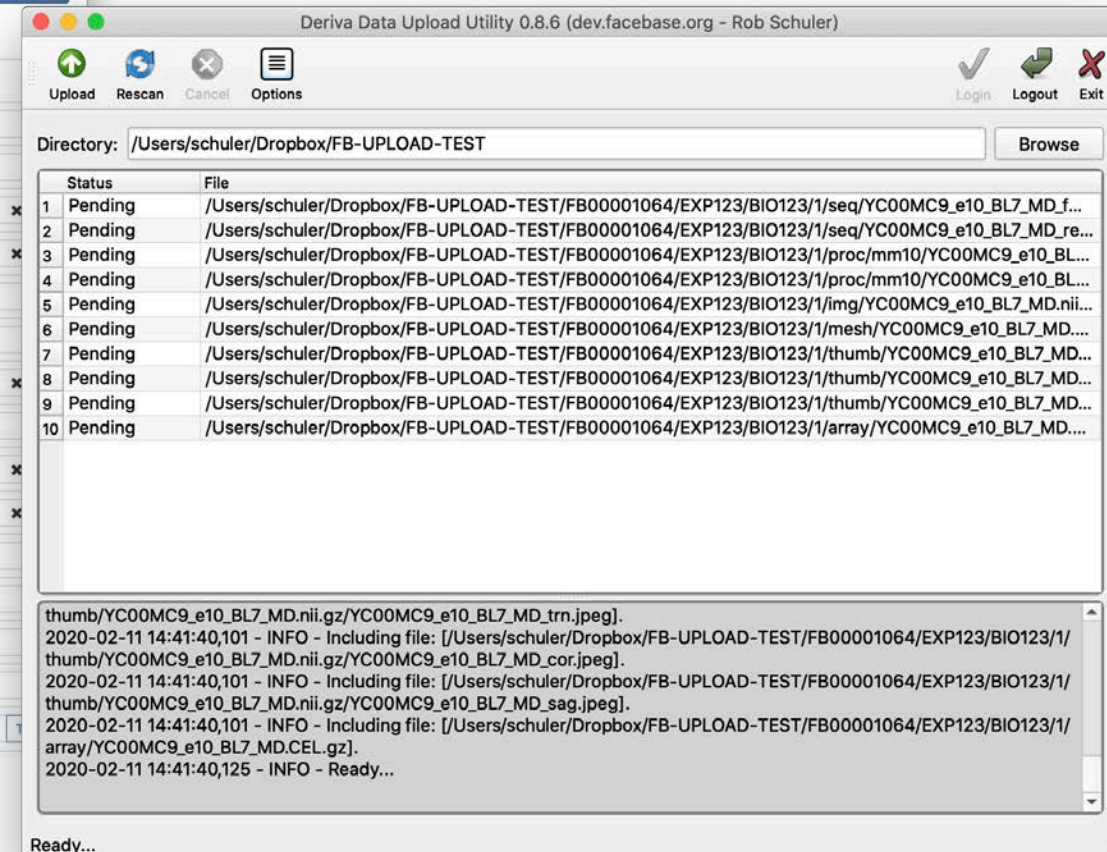
FaceBase Data Browser Submit Resources About News Help schuler@globusid.org

Edit Biosample

* indicates required field

Record Number	1	2
* Dataset	Select a value	Select a value
Local Identifier	3XhSox9-E125-S5_pHsp68-lacZ-tdTom_E9.5_I-42	hSox9_145mb_Enh-pHsp68-lacZ-tdTomato_E9.5_I-42
* Species	Mus musculus	Mus musculus
* Specimen	whole organism preparation	whole organism preparation
Gene	Select a value	Select a value
Genotype	Select a value	Select a value
Strain	FVB	FVB
Mutation	Select a value	Select a value
Stage	E9.5	E9.5
Anatomy	head	head
Origin	Select a value	Select a value
Phenotype	Select a value	Select a value
Gender	Select a value	Select a value
Litter		
Collection Date	YYYY-MM-DD Today	YYYY-MM-DD

(2) Graphical desktop applications



(1) Online forms with multi-record entry and edit

(3) Command-line, Python and Web services interfaces for custom data submission workflows (not pictured)

Researchers have submitted their own datasets with 100s to 1000s of files, usually in a few days

Quality Control Dashboards

- Automated Quality Control Rules to check each dataset
- QC issues are displayed on project pages (visible only to contributors)

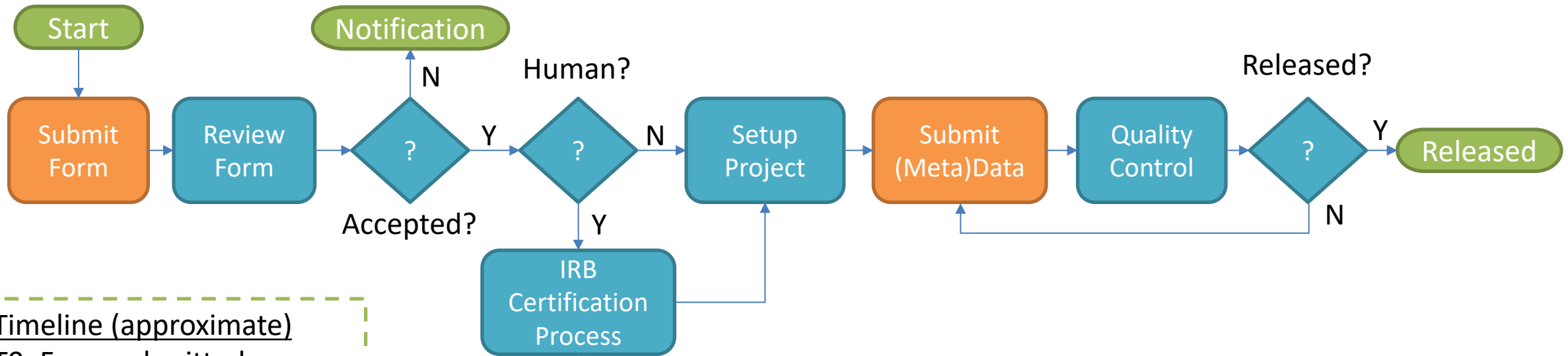
Automated QC Issue Flagging

The screenshot shows the FaceBase Data Browser interface. The top section displays project information for 'Chai: Integrated research of craniofacial morphogenesis'. Below this, a table lists datasets with their respective quality control issues. A red arrow points from the 'Automated QC Issue Flagging' text to the 'Quality Control Issues' column in the table.

Release Date	Quality Control Issues
	Experiment missing at least one replicate
	Experiment missing at least one replicate
	Missing dataset stage tag, Experiment missing at least one replicate
2019-04-15	Replicate(s) missing data files
2019-04-15	Missing dataset anatomy tag, Replicate(s) missing data files
2019-04-15	Replicate(s) missing data files
2019-04-15	Replicate(s) missing data files

Quality Control "Dashboard" Integrated into Project Pages

Streamlined Process for Data Submission



Timeline (approximate)

- T0: Form submitted
- T+2 weeks: Review decision
- T+3 weeks: Project setup
- T+5 weeks: Submit data*
- T+6 weeks: QC review
- * Based on user averages

IRB Certification Process

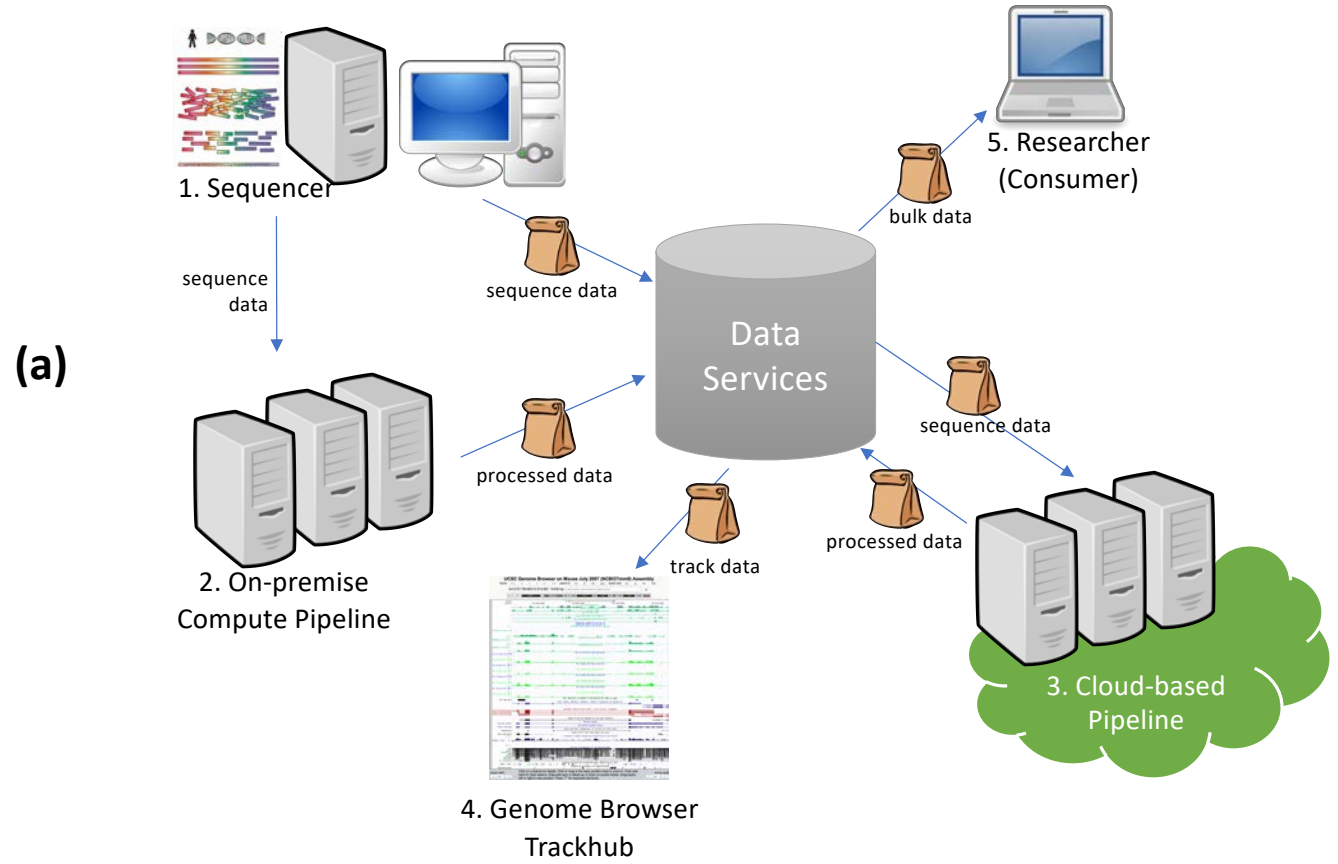
- Individual level data classified as human subjects
- Requires USC certification of your IRB decision
- Tracks are not considered restricted data
- Timeline Varies



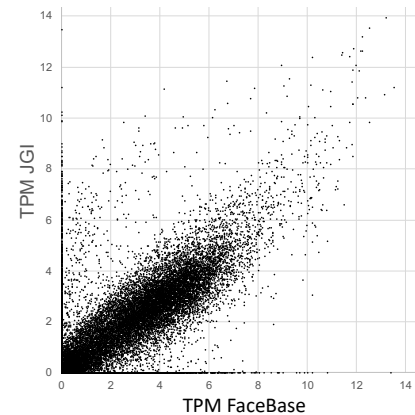
A Test of the Reproducibility of FaceBase Data

Evaluation: 3 labs, 13 datasets, on 192 RNA-Seq or ChIP-Seq experiments (1000+ data files)

- a) Uniform processing pipeline
- b) Output strongly correlated with researcher's results



(b)



RNA-seq log2 counts

		LBNL ChIP-seq peaks															
		H3K27ac				H3K27me3				H3K4me1							
		lnp	md	mnp	mx	lnp	md	mnp	mx	lnp	md	mnp	mx				
FaceBase ChIP-seq peaks	H3K27ac	lnp	1519	1370	1409	1429	7	11	16	11	279	226	272	260			
		md	1461	1578	1417	1421	18	6	22	6	304	304	308	313			
		mnp	1485	1371	1514	1393	10	10	14	11	286	272	320	295			
		mx	1499	1442	1436	1637	15	6	17	2	262	259	279	290			
	H3K27me3	lnp	3	35	14	24	1859	1723	1802	1696	47	42	58	45			
		md	12	2	17	6	1762	1868	1869	1809	60	51	69	33			
		mnp	16	21	20	20	1747	1794	1859	1768	63	62	70	35			
		mx	7	20	11	4	1816	1816	1877	1851	45	41	49	25			
	H3K4me1	lnp	425	419	442	351	50	60	71	62	1058	688	848	845			
		md	369	403	383	340	78	84	94	79	669	1009	763	767			
		mnp	432	416	461	376	47	59	66	56	850	771	1125	902			
		mx	480	476	499	460	21	20	25	21	889	869	961	1189			

ChIP-seq peaks (overlap top-2000)

Thank You

- **Co-PIs:** Carl Kesselman; Yang Chai
- **Core Team:** Rob Schuler (CS & Technical); Bridget Samuels (Biocuration); Alejandro Bugacov (Data science); Cris Williams (Communications); Joe Hacia (Bioinformatics); Thach Vu Ho (Data curation)
- **Website:** www.facebase.org
- **Sponsor:** NIH / NIDCR (U01DE028729)