

Human Genomics Analysis Interface

Mary L. Marazita

Elizabeth Leslie, Eleanor Feingold, Harry Hochheiser

University of Pittsburgh

Center for Craniofacial and Dental Genetics



FaceBase Annual Meeting
May 2-3, 2016



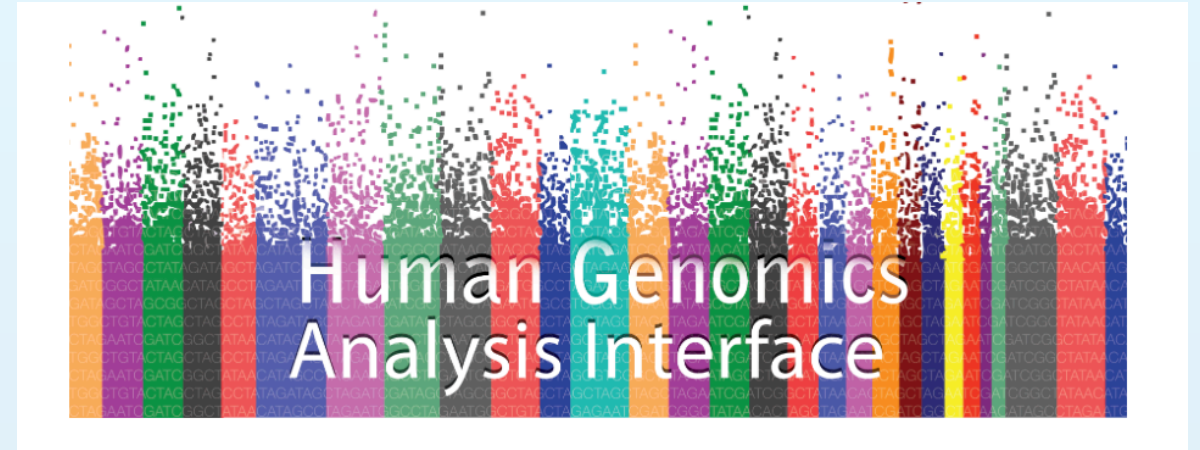
GOALS

- Develop software interface for Human Genomics Results
 - No access to individual level data
 - Secure hardware
- Identify appropriate craniofacial genomics projects
 - From craniofacial research community, dbGaP, publications
- Create and display results from genomics projects
 - E.g. Manhattan plots, LocusZoom



UPDATES

- Hardware established
- Basic software created
- WebSite created and updated
- Multiple projects identified
 - Orofacial Clefting, Facial Variation, Dental
- Project pages created
- Results databases and interface created



APPROACH

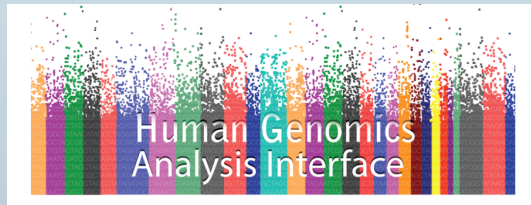
- User friendly, save user time
- Pre-calculate results databases (e.g. p-values at SNPs)
 - For all relevant subsets: by ethnicity, by genotyped versus imputed panels, by phenotype, by environmental factors
 - Customized: with PIs (e.g. published results)
- Prepare Manhattan plot files
- Pre-run typical LocusZoom requests

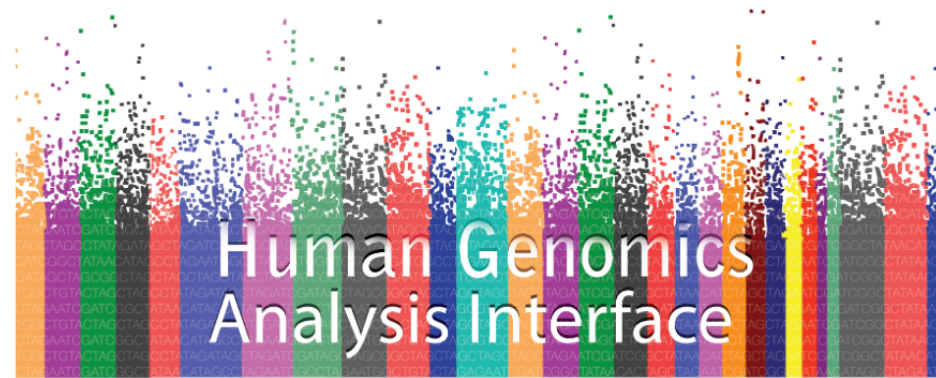


WEBSITE

FaceBase.sdmgenetics.pitt.edu

- Link also available on the FaceBase.org Home page, and our Project page





Human Genomics Analysis Interface



The Human Genomics Analysis Interface is a project of the University of Pittsburgh Center for Craniofacial and Dental Genetics (Director: Mary L. Marazita), and is part of the [FaceBase](#) consortium. This project is possible from funding by NIDCR, Grant #: [U01-DE024425](#).

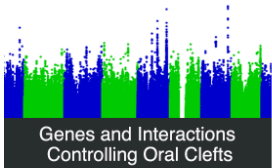
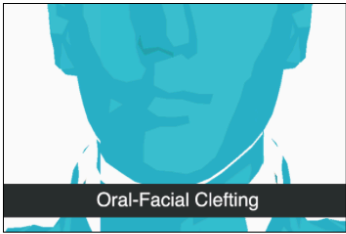
There are now several large human genomics databases relevant to craniofacial research, including multiple databases funded, in part, under the FaceBase consortium. Direct access to the individual level data from such databases can be cumbersome; therefore the Human Genomics Analysis Interface seeks to make analyses of pertinent genomics data available to craniofacial users without needing access to the individual level data. This project will provide multiple avenues to explore existing data.



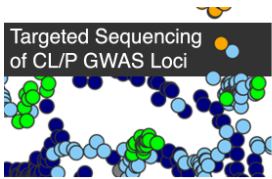
© 2014-2016 University of Pittsburgh.

This project is made possible by funding from NIDCR.
Grant #: U01-DE024425





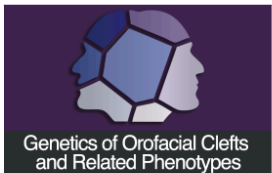
[International Consortium to Identify Genes and Interactions Controlling Oral Clefts- Genome Wide Association Study](#)
dbGaP Study Accession: phs000094.v1.p1
Principal Investigator: Terri H. Beaty, Johns Hopkins School of Public Health, Baltimore, MD, USA



[Targeted Sequencing of CL/P GWAS Loci](#)
dbGaP Study Accession: phs000625.v1.p1
Principal Investigators: Jeffrey C. Murray, University of Iowa, Iowa City, IA, USA
Mary L. Marazita, University of Pittsburgh, Pittsburgh, PA, USA
Richard K. Wilson, Washington University School of Medicine, St. Louis, MO, USA
George Weinstock, Jackson Laboratory for Genomic Medicine, Farmington, CT, USA



[GWAS of OFCs in Guatemala](#)
dbGaP Study Accession: phs000440.v1.p1
Principal Investigator: Mary Marazita, University of Pittsburgh School of Dental Medicine, Pittsburgh, PA, USA

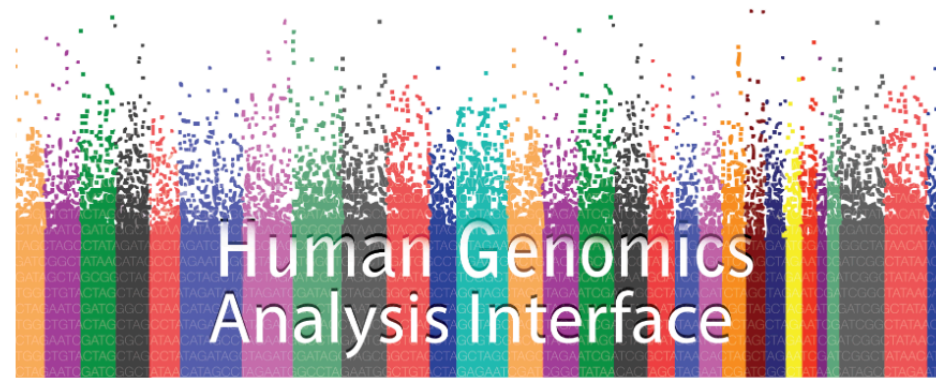


[Genetics of Orofacial Clefts and Related Phenotypes](#)
dbGaP Study Accession: phs000774.v1.p1
Principal Investigator: Mary Marazita, University of Pittsburgh School of Dental Medicine, Pittsburgh, PA, USA

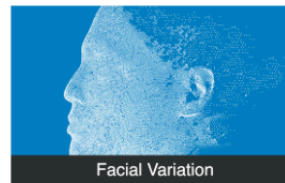


© 2014-2016 University of Pittsburgh.
This project is made possible by funding from NIDCR.
Grant #: U01-DE024425





Human Genomics Analysis Interface



The Human Genomics Analysis Interface is a project of the University of Pittsburgh Center for Craniofacial and Dental Genetics (Director: Mary L. Marazita), and is part of the [FaceBase](#) consortium. This project is possible from funding by NIDCR, Grant #: [U01-DE024425](#).

There are now several large human genomics databases relevant to craniofacial research, including multiple databases funded, in part, under the FaceBase consortium. Direct access to the individual level data from such databases can be cumbersome; therefore the Human Genomics Analysis Interface seeks to make analyses of pertinent genomics data available to craniofacial users without needing access to the individual level data. This project will provide multiple avenues to explore existing data.



© 2014-2016 University of Pittsburgh.

This project is made possible by funding from NIDCR.
Grant #: U01-DE024425

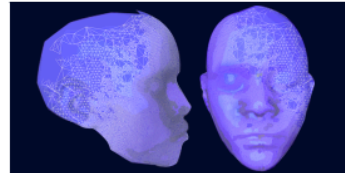




[3D Facial Norms](#)

dbGaP Study Accession:

Principal Investigator: Seth Weinberg. University of Pittsburgh School of Dental Medicine, Pittsburgh, PA, USA



[Genetic Determinants of Orofacial Shape and Relationship to Cleft Lip/Palate](#)

dbGaP Study Accession: phs000622.v1.p1

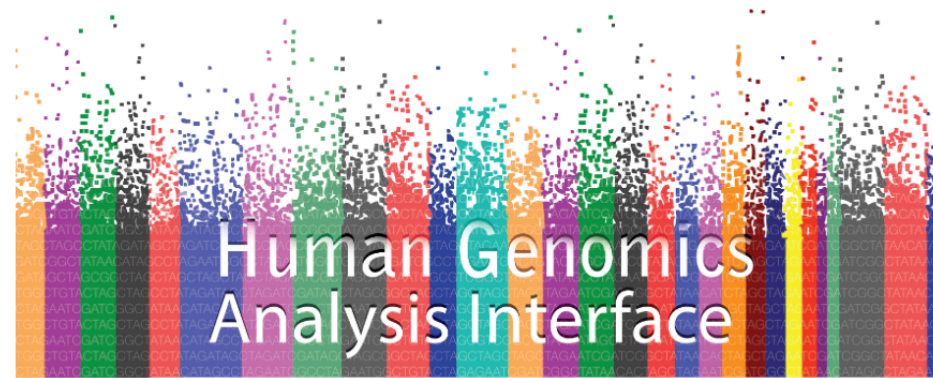
Principal Investigator: Richard Spritz, MD. University of Colorado School of Medicine, Aurora, CO USA



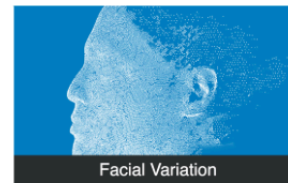
© 2014-2016 University of Pittsburgh.

This project is made possible by funding from NIDCR.
Grant #: U01-DE024425





Human Genomics Analysis Interface



The Human Genomics Analysis Interface is a project of the University of Pittsburgh Center for Craniofacial and Dental Genetics (Director: Mary L. Marazita), and is part of the [FaceBase](#) consortium. This project is possible from funding by NIDCR, Grant #: [U01-DE024425](#).

There are now several large human genomics databases relevant to craniofacial research, including multiple databases funded, in part, under the FaceBase consortium. Direct access to the individual level data from such databases can be cumbersome; therefore the Human Genomics Analysis Interface seeks to make analyses of pertinent genomics data available to craniofacial users without needing access to the individual level data. This project will provide multiple avenues to explore existing data.



© 2014-2016 University of Pittsburgh.

This project is made possible by funding from NIDCR.
Grant #: U01-DE024425





[Childhood Caries](#)

dbGap Study Accession: phs000095.v3.p1

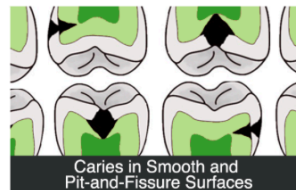
Principal Investigator: Mary Marazita, University of Pittsburgh School of Dental Medicine, Pittsburgh, PA, USA



[Caries in Permanent Dentition](#)

dbGap Study Accession: phs000095.v3.p1

Principal Investigator: Mary Marazita, University of Pittsburgh School of Dental Medicine, Pittsburgh, PA, USA



[Caries in Smooth and Pit-and-Fissure Surfaces](#)

dbGap Study Accession: phs000095.v3.p1

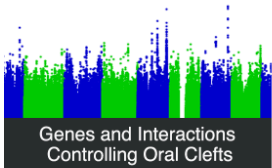
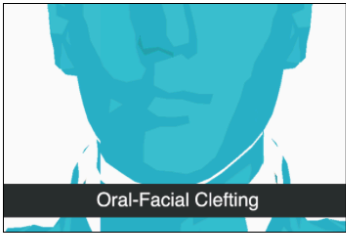
Principal Investigator: Mary Marazita, University of Pittsburgh School of Dental Medicine, Pittsburgh, PA, USA



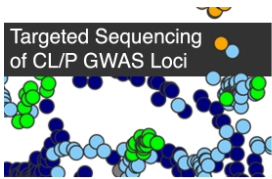
© 2014-2016 University of Pittsburgh.

This project is made possible by funding from NIDCR.
Grant #: U01-DE024425





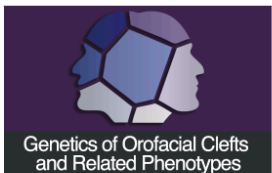
[International Consortium to Identify Genes and Interactions Controlling Oral Clefts- Genome Wide Association Study](#)
dbGaP Study Accession: phs000094.v1.p1
Principal Investigator: Terri H. Beaty, Johns Hopkins School of Public Health, Baltimore, MD, USA



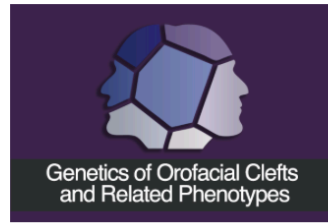
[Targeted Sequencing of CL/P GWAS Loci](#)
dbGaP Study Accession: phs000625.v1.p1
Principal Investigators: Jeffrey C. Murray, University of Iowa, Iowa City, IA, USA
Mary L. Marazita, University of Pittsburgh, Pittsburgh, PA, USA
Richard K. Wilson, Washington University School of Medicine, St. Louis, MO, USA
George Weinstock, Jackson Laboratory for Genomic Medicine, Farmington, CT, USA



[GWAS of OFCs in Guatemala](#)
dbGaP Study Accession: phs000440.v1.p1
Principal Investigator: Mary Marazita, University of Pittsburgh School of Dental Medicine, Pittsburgh, PA, USA



[Genetics of Orofacial Clefts and Related Phenotypes](#)
dbGaP Study Accession: phs000774.v1.p1
Principal Investigator: Mary Marazita, University of Pittsburgh School of Dental Medicine, Pittsburgh, PA, USA



- Project Description
- Methods
- Descriptive Statistics
- Explore Project Data
- Publications

dbGaP Study Accession: [phs000774.v1.p1](#)

Principal Investigator: Mary L. Marazita, University of Pittsburgh, Pittsburgh, PA, USA

Funding Sources:

X01-HG00784 Genetics of Orofacial Clefts and Related Phenotypes. National Human Genome Research Institute, CIDR Genotyping, National Institutes of Health, Bethesda, MD, USA

R01-DE-016148, Extending the Phenotype of Nonsyndromic Orofacial Clefts. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R01-DE-012472, HOMOZYGOSITY MAPPING OF ORAL-FACIAL CLEFTS IN TURKEY. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R01-DE-009886, GENE MAPPING STUDIES OF ORAL/FACIAL CLEFTS - CHINA/INDIA. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

P50-DE-016215, CRANIOFACIAL ANOMALIES RESEARCH CENTER. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

K99-DE-022378, GENETIC STUDIES OF NONSYNDROMIC CLEFTS IN POPULATIONS OF AFRICAN DESCENT. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R37-DE-008559, MOLECULAR GENETIC EPIDEMIOLOGY OF CLEFT LIP AND PALATE. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R01-DE-014667, CLEFT LIP GENETICS: A MULTI CENTER INTERNATIONAL CONSORTIUM. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R01-DD-000295, HEALTH OUTCOMES AND IMPROVED PHENOTYPIC CHARACTERIZATION OF CLEFT LIP AND PALATE. National Center on Birth Defects and Developmental Disabilities, Centers for Disease Control and Prevention, Atlanta, GA, USA

R21-DE-016930, PLANNING INTERNATIONAL OROFACIAL CLEFT GENETIC STUDIES. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

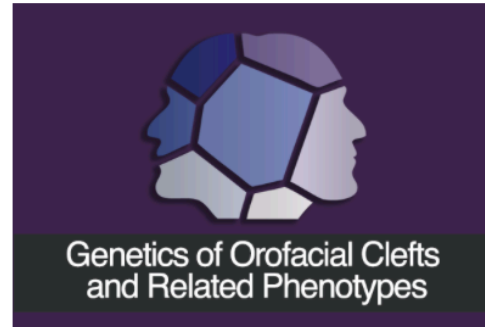
The current study is the culmination of many years of ongoing collaborations among the study investigators (for complete list of investigators, see [dbGaP](#)). Molecular genetic studies of orofacial clefts (OFC) by members of this research consortium began in the 1980's, followed by genome-wide linkage studies in the 1990's-early 2000's, and then genome-wide association studies in the 2010's. A rich phenotyping approach was first added to the consortium in 1999 as a project led by Dr. Mary Marazita under a center grant headed by Dr. Jeff Murray. Additional sites were added throughout the 2000's, until multiple populations and a large number of individuals (~12,000) comprise the current study population for genetic studies of OFCs and related subclinical phenotypes—subtle features believed to represent mild manifestations of the same underlying genetic susceptibility responsible for OFCs. As such, their inclusion in case-control and family-based genetic studies can help to clarify and refine the relationship between genotype and phenotype. Information on how to obtain individual level data is available on [dbGaP](#).



© 2014-2016 University of Pittsburgh.

This project is made possible by funding from NIDCR.
Grant #: U01-DE024425





- Project Description
- Methods
- Descriptive Statistics
- Explore Project Data
- Publications

Methods

Genotyping and quality control methods are available on dbGaP and are briefly described in Leslie et al. [“A genome-wide association study of nonsyndromic cleft palate identifies an etiologic missense variant in GRHL3”](#). Am J Hum Genet. 2016; and Leslie et al. [“A multi-ethnic genome-wide association study identifies novel loci for nonsyndromic cleft lip with or without cleft palate on 2p24.2, 17q23 and 19q31”](#). Hum Mol Genet. 2016.

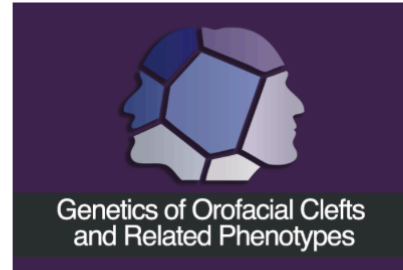
Briefly, GWAS scans were conducted in two subsets: independent cases and controls, and an independent set of case-parent trios. Statistical analysis was performed in PLINK using logistic regression while adjusting for principal components of ancestry and the transmission disequilibrium test. Association results for the two GWAS scans were combined by weighted odds ratio meta-analysis to estimate study-wide association results. Analyses were completed in the full multiethnic sample or stratified by genetically defined subpopulations.



© 2014-2016 University of Pittsburgh.

This project is made possible by funding from NIDCR.
Grant #: U01-DE024425



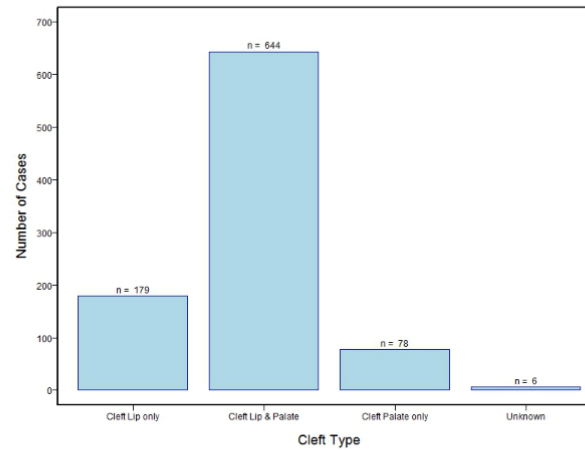


- Project Description
- Methods
- Descriptive Statistics
- Explore Project Data
- Publications

Frequency Counts for Study Variables

Please select option:

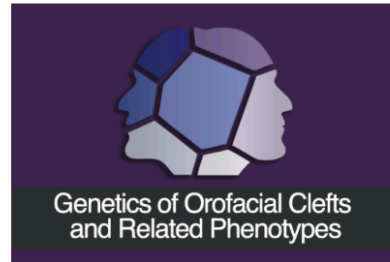
Case Clefting Status



© 2014-2016 University of Pittsburgh.

This project is made possible by funding from NIDCR.
Grant #: U01-DE024425





- Project Description
- Methods
- Descriptive Statistics
- Explore Project Data**
- Publications

Explore Project Data

The samples for this study come from three primary ancestry groups: Europeans (from the United States, Denmark, Hungary, Spain, and Turkey), Asians (from China, the Philippines, and India), Central/South Americans (from Puerto Rico, Guatemala, Colombia, and Argentina), and Africans (from Nigeria and Ethiopia). The subjects have been assessed for various cleft types and related subclinical phenotypes. Our results database contains the GWAS results from this study of OFCs and can be explored as one study or subset into separate groups.

Custom Plots

Please select one option from each of the categories. You must select one option from each available category to be able to submit the request.

Genotype Data <input checked="" type="radio"/> Genotyped <input type="radio"/> Imputed	Phenotype <input checked="" type="radio"/> Cleft Palate (CP) <input type="radio"/> Cleft Lip With Or Without Cleft Palate (CL/P)
Ancestry Groups <input checked="" type="radio"/> All <input type="radio"/> Asian <input type="radio"/> European <input type="radio"/> Central/South Americans	Analysis Group <input checked="" type="radio"/> Trios <input type="radio"/> Case-Control <input type="radio"/> Meta-analysis
	Output <input checked="" type="radio"/> Manhattan Plot <input type="radio"/> Region/Locus

Submit Reset



Project
Description

Methods

Descriptive
StatisticsExplore
Project Data

Publications

Explore Project Data

The samples for this study come from two primary ancestry groups: Asians (from China, the Philippines, Singapore, Korea, and Taiwan) and Europeans (from the United States, Norway, and Denmark). The subjects have been assessed for various cleft types and reported on first trimester environmental exposures. Our results database contains the GWAS results from this study of CL/P and can be explored as one study or subsetted into separate groups.

View published peaks

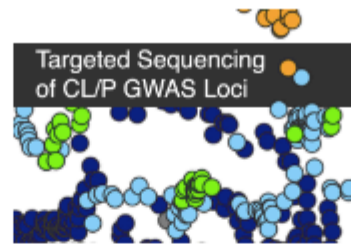
These are peaks that have been published in [Nature Genetics](#). They are for all ancestries, all cleft types, and for imputed data.

Custom Plots

Please select one option from each of the categories. You must select one option from each available category to be able to submit the request.

***Note:** There is no Imputed data available for hg18.

Genotype Data <input checked="" type="radio"/> Genotyped <input type="radio"/> Imputed	Environmental Factors <small>*"Yes" selects only subjects who responded "yes" to an exposure during the first trimester. "No" selects only subjects who responded "no". If no subsetting by environmental factors is desired, please select "none."*</small> Smoking <input type="radio"/> Yes <input type="radio"/> No Alcohol <input type="radio"/> Yes <input type="radio"/> No Vitamin Use <input type="radio"/> Yes <input type="radio"/> No <input checked="" type="radio"/> None
Genome Build <input checked="" type="radio"/> hg18 <input type="radio"/> hg19	Output <input checked="" type="radio"/> Manhattan Plot <input type="radio"/> Region/Locus
Phenotype <input checked="" type="radio"/> All <input type="radio"/> Cleft Lip (CL) <input type="radio"/> Cleft Lip With Cleft Palate (CLP) <input type="radio"/> Cleft Palate (CP) <input type="radio"/> Cleft Lip With Or Without Cleft Palate (CL/P) <input type="radio"/> Cleft Palate With Or Without Cleft Lip (CP/L)	Ancestry Groups <input checked="" type="radio"/> All <input type="radio"/> Asian <input type="radio"/> European
<input type="button" value="Submit"/> <input type="button" value="Reset"/>	



Project Description | Methods | Descriptive Statistics | **Explore Project Data** | Publications

Explore Project Data

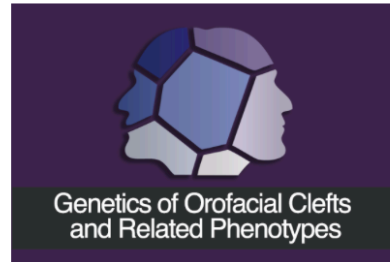
This project involved targeted sequencing of GWAS loci for CL/P. Thirteen regions were sequenced. The samples for this study come from two primary ancestry groups: Asians (from China and the Philippines) and Europeans (from several sites in the United States). Our results database contains the association results for each of the thirteen sequenced regions and can be explored in the combined sample or in each ancestry group separately.

Custom Plots

Please select one option from each of the categories. You must select one option from each available category to be able to submit the request.

<p>Ancestry Groups</p> <ul style="list-style-type: none"><input type="radio"/> All<input type="radio"/> Asian<input type="radio"/> European	<p>Locus</p> <ul style="list-style-type: none"><input type="radio"/> 1p36 (PAX7)<input type="radio"/> 1p22 (ARHGAP29)<input type="radio"/> 1q32 (IRF6)<input type="radio"/> 4p16 (MSX1)<input type="radio"/> 8q24<input type="radio"/> 9q22 (FOXE1)<input type="radio"/> 9q22 (PTCH1)<input type="radio"/> 10q25 (VAX1)<input type="radio"/> 10q26 (FGFR2)<input type="radio"/> 14q22 (BMP4)<input type="radio"/> 17p13 (NTN1)<input type="radio"/> 17q22 (NOG)<input type="radio"/> 20q12 (MAFB)
---	---





- Project Description
- Methods
- Descriptive Statistics
- Explore Project Data**
- Publications

Explore Project Data

The samples for this study come from three primary ancestry groups: Europeans (from the United States, Denmark, Hungary, Spain, and Turkey), Asians (from China, the Philippines, and India), Central/South Americans (from Puerto Rico, Guatemala, Colombia, and Argentina), and Africans (from Nigeria and Ethiopia). The subjects have been assessed for various cleft types and related subclinical phenotypes. Our results database contains the GWAS results from this study of OFCs and can be explored as one study or subset into separate groups.

Custom Plots

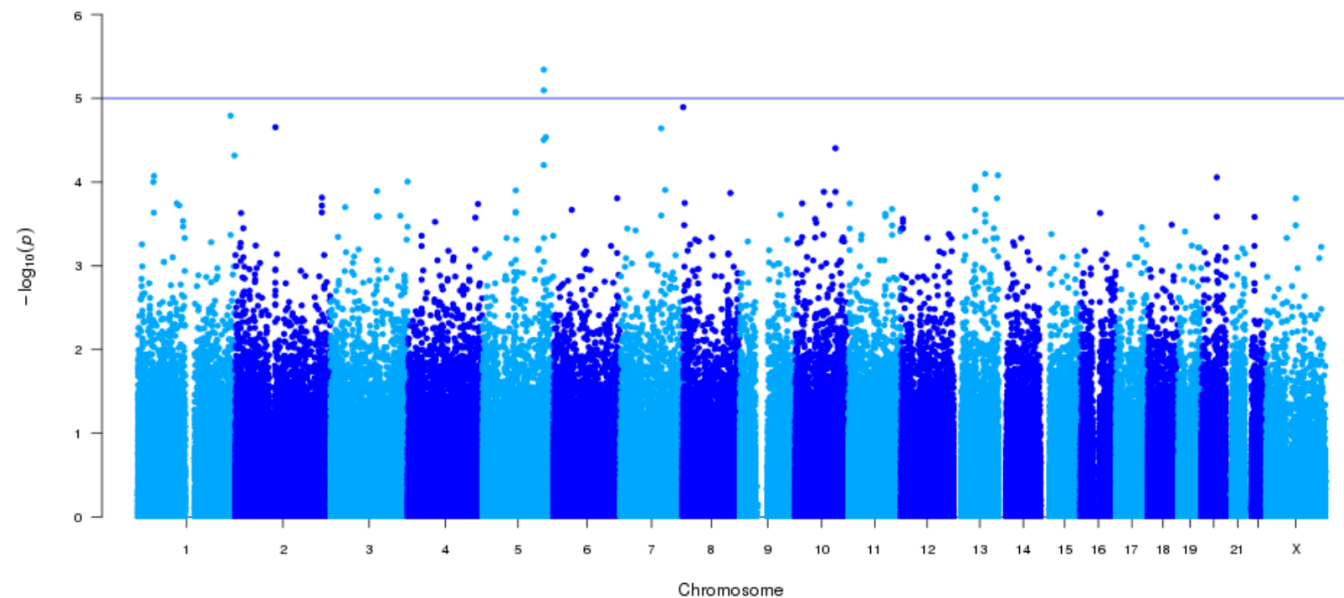
Please select one option from each of the categories. You must select one option from each available category to be able to submit the request.

Genotype Data <input checked="" type="radio"/> Genotyped <input type="radio"/> Imputed	Phenotype <input checked="" type="radio"/> Cleft Palate (CP) <input type="radio"/> Cleft Lip With Or Without Cleft Palate (CL/P)
Ancestry Groups <input checked="" type="radio"/> All <input type="radio"/> Asian <input type="radio"/> European <input type="radio"/> Central/South Americans	Analysis Group <input checked="" type="radio"/> Trios <input type="radio"/> Case-Control <input type="radio"/> Meta-analysis
	Output <input checked="" type="radio"/> Manhattan Plot <input type="radio"/> Region/Locus

Submit Reset



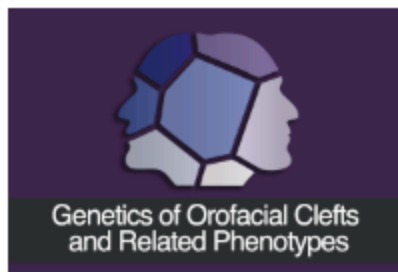
OFC GWAS (genotyped) - TDT - ALLPOPS - CP



© 2014-2016 University of Pittsburgh.

This project is made possible by funding from NIDCR.
Grant #: U01-DE024425



Project
Description

Methods

Descriptive
StatisticsExplore
Project Data

Publications

Explore Project Data

The samples for this study come from three primary ancestry groups: Europeans (from the United States, Denmark, Hungary, Spain, and Turkey), Asians (from China, the Philippines, and India), Central/South Americans (from Puerto Rico, Guatemala, Colombia, and Argentina), and Africans (from Nigeria and Ethiopia). The subjects have been assessed for various cleft types and related subclinical phenotypes. Our results database contains the GWAS results from this study of OFCs and can be explored as one study or subset into separate groups.

Custom Plots

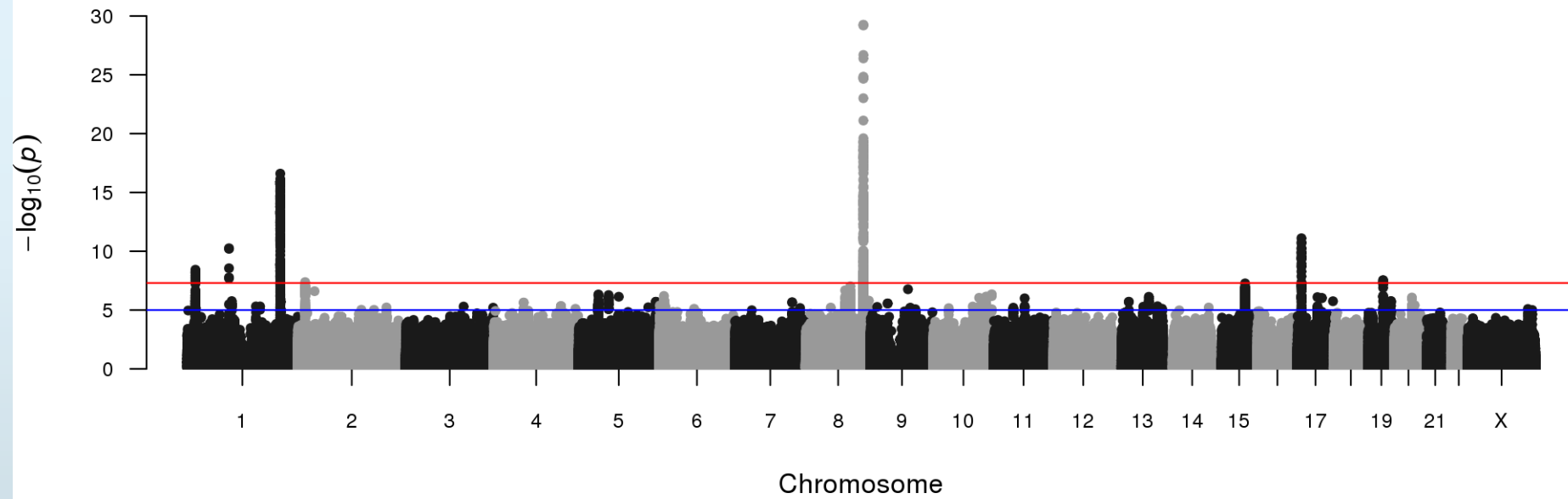
Please select one option from each of the categories. You must select one option from each available category to be able to submit the request.

Genotype Data <input type="radio"/> Genotyped <input checked="" type="radio"/> Imputed	Phenotype <input type="radio"/> Cleft Palate (CP) <input checked="" type="radio"/> Cleft Lip With Or Without Cleft Palate (CL/P)
Ancestry Groups <input checked="" type="radio"/> All <input type="radio"/> Asian <input type="radio"/> European <input type="radio"/> Central/South Americans	Analysis Group <input type="radio"/> Trios <input type="radio"/> Case-Control <input checked="" type="radio"/> Meta-analysis
	Output <input checked="" type="radio"/> Manhattan Plot <input type="radio"/> Region/Locus

Submit

Reset

OFC GWAS (imputed) - Meta Analysis - ALLPOPS - CLCLP



Project
Description

Methods

Descriptive
StatisticsExplore
Project Data

Publications

Explore Project Data

The samples for this study come from three primary ancestry groups: Europeans (from the United States, Denmark, Hungary, Spain, and Turkey), Asians (from China, the Philippines, and India), Central/South Americans (from Puerto Rico, Guatemala, Colombia, and Argentina), and Africans (from Nigeria and Ethiopia). The subjects have been assessed for various cleft types and related subclinical phenotypes. Our results database contains the GWAS results from this study of OFCs and can be explored as one study or subset into separate groups.

Custom Plots

Please select one option from each of the categories. You must select one option from each available category to be able to submit the request.

Genotype Data <input checked="" type="radio"/> Genotyped <input type="radio"/> Imputed	Phenotype <input checked="" type="radio"/> Cleft Palate (CP) <input type="radio"/> Cleft Lip With Or Without Cleft Palate (CL/P)
Ancestry Groups <input type="radio"/> All <input checked="" type="radio"/> Asian <input type="radio"/> European <input type="radio"/> Central/South Americans	Analysis Group <input checked="" type="radio"/> Trios <input type="radio"/> Case-Control <input type="radio"/> Meta-analysis
	Output <input checked="" type="radio"/> Manhattan Plot <input type="radio"/> Region/Locus

Submit

Reset

Project Description

Methods

Descriptive Statistics

Explore Project Data

Publications

Explore Project Data

The samples for this study come from three primary ancestry groups: Europeans (from the United States, Denmark, Hungary, Spain, and Turkey), Asians (from China, the Philippines, and India), Central/South Americans (from Puerto Rico, Guatemala, Colombia, and Argentina), and Africans (from Nigeria and Ethiopia). The subjects have been assessed for various cleft types and related subclinical phenotypes. Our results database contains the GWAS results from this study of OFCs and can be explored as one study or subset into separate groups.

Custom Plots

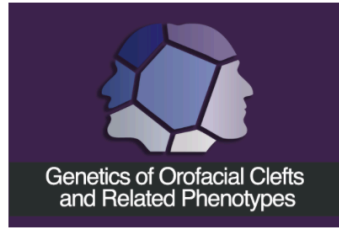
Please select one option from each of the categories. You must select one option from each available category to be able to submit the request.

Genotype Data <input checked="" type="radio"/> Genotyped <input type="radio"/> Imputed	Phenotype <input checked="" type="radio"/> Cleft Palate (CP) <input type="radio"/> Cleft Lip With Or Without Cleft Palate (CL/P)
Ancestry Groups <input type="radio"/> All <input checked="" type="radio"/> Asian <input type="radio"/> European <input type="radio"/> Central/South Americans	Analysis Group <input checked="" type="radio"/> Trios <input type="radio"/> Case-Control <input type="radio"/> Meta-analysis
Output <input checked="" type="radio"/> Manhattan Plot <input type="radio"/> Region/Locus	

The selected values represent a low number of trios so a graph was not generated. Please change an option and try again.

Close





- Project Description
- Methods
- Descriptive Statistics
- Explore Project Data
- Publications

Explore Project Data

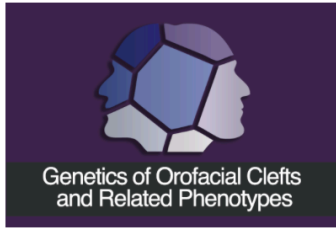
The samples for this study come from three primary ancestry groups: Europeans (from the United States, Denmark, Hungary, Spain, and Turkey), Asians (from China, the Philippines, and India), Central/South Americans (from Puerto Rico, Guatemala, Colombia, and Argentina), and Africans (from Nigeria and Ethiopia). The subjects have been assessed for various cleft types and related subclinical phenotypes. Our results database contains the GWAS results from this study of OFCs and can be explored as one study or subset into separate groups.

Custom Plots

Please select one option from each of the categories. You must select one option from each available category to be able to submit the request.

Genotype Data <input checked="" type="radio"/> Genotyped <input type="radio"/> Imputed	Phenotype <input checked="" type="radio"/> Cleft Palate (CP) <input type="radio"/> Cleft Lip With Or Without Cleft Palate (CL/P)
Ancestry Groups <input checked="" type="radio"/> All <input type="radio"/> Asian <input type="radio"/> European <input type="radio"/> Central/South Americans	Analysis Group <input checked="" type="radio"/> Trios <input type="radio"/> Case-Control <input type="radio"/> Meta-analysis
	Output <input type="radio"/> Manhattan Plot <input checked="" type="radio"/> Region/Locus Please Select an option Explore custom regions <input checked="" type="radio"/> Gene <input type="radio"/> SNP <input type="radio"/> Ch: BP-BP <input type="text" value="IRF6"/>





- Project Description
- Methods
- Descriptive Statistics
- Explore Project Data
- Publications

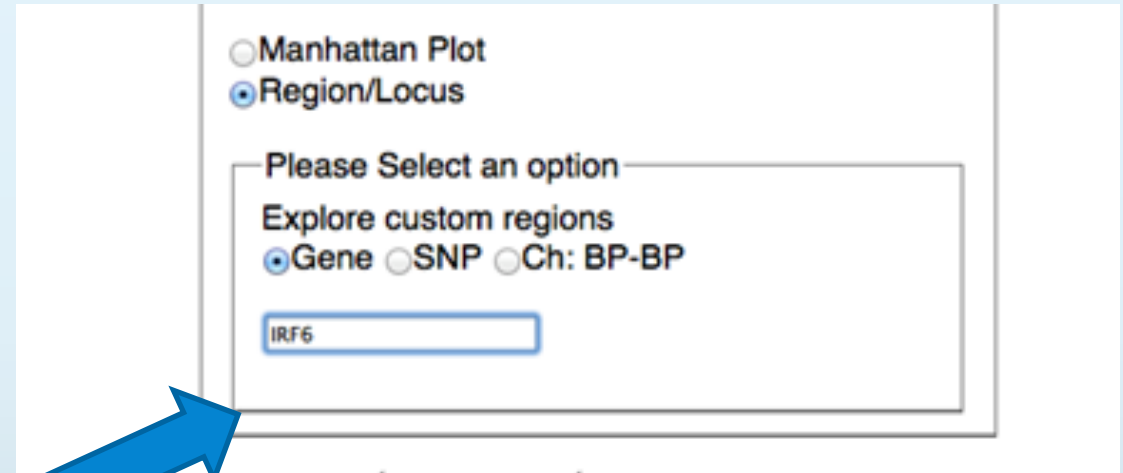
Explore Project Data

The samples for this study come from three primary ancestry groups: Europeans (from the United States, Denmark, Hungary, Spain, and Turkey), Asians (from China, the Philippines, and India), Central/South Americans (from Puerto Rico, Guatemala, Colombia, and Argentina), and Africans (from Nigeria and Ethiopia). The subjects have been assessed for various cleft types and related subclinical phenotypes. Our results database contains the GWAS results from this study of OFCs and can be explored as one study or subset into separate groups.

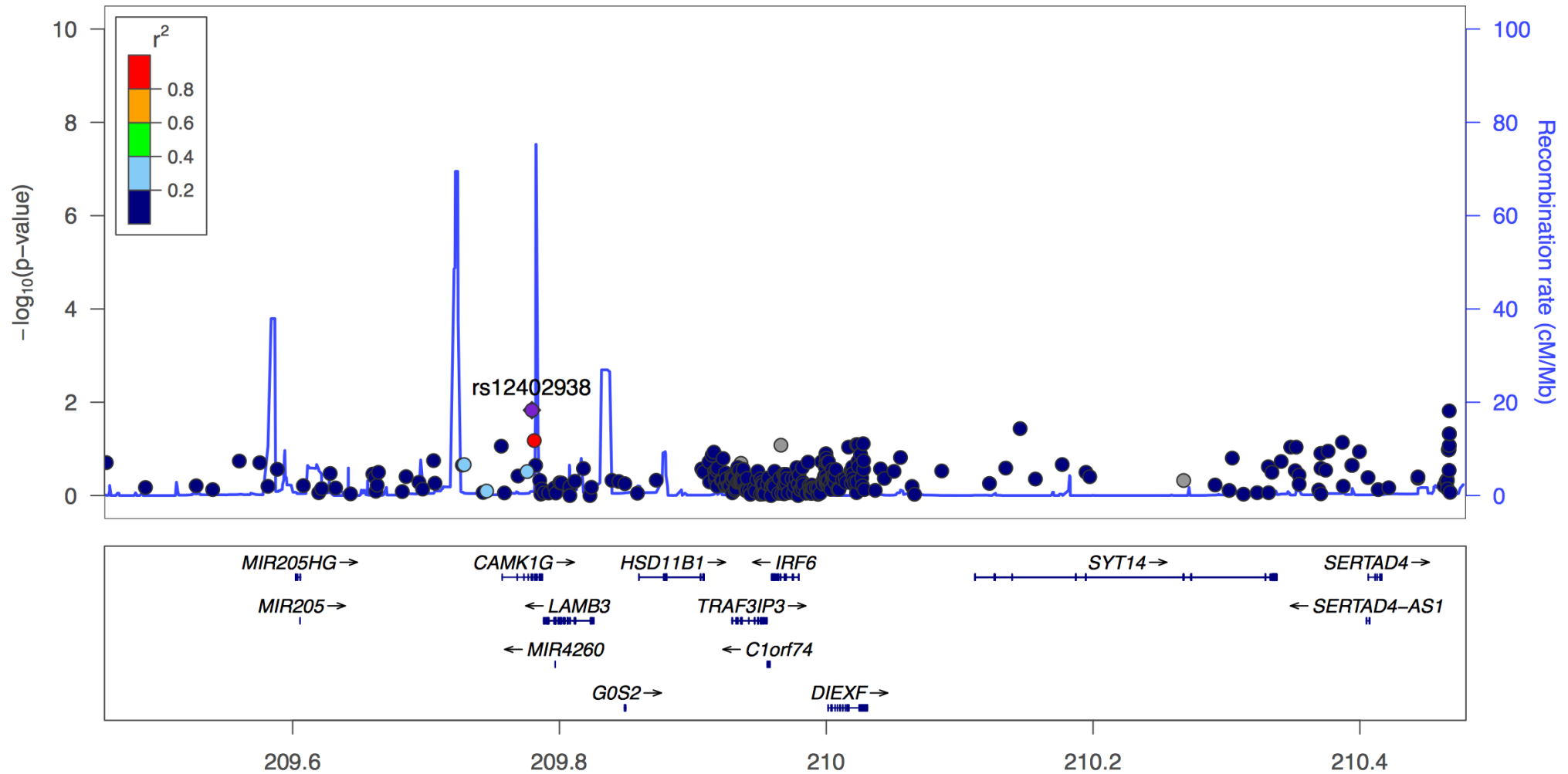
Custom Plots

Please select one option from each of the categories. You must select one option from each available category to be able to submit the request.

Genotype Data <input checked="" type="radio"/> Genotyped <input type="radio"/> Imputed	Phenotype <input checked="" type="radio"/> Cleft Palate (CP) <input type="radio"/> Cleft Lip With Or Without Cleft Palate (CL/P)
Ancestry Groups <input checked="" type="radio"/> All <input type="radio"/> Asian <input type="radio"/> European <input type="radio"/> Central/South Americans	Analysis Group <input checked="" type="radio"/> Trios <input type="radio"/> Case-Control <input type="radio"/> Meta-analysis
	Output <input type="radio"/> Manhattan Plot <input checked="" type="radio"/> Region/Locus Please Select an option Explore custom regions <input checked="" type="radio"/> Gene <input type="radio"/> SNP <input type="radio"/> Ch: BP-BP <input type="text" value="IRF6"/>



Geno HG19, All Ancestry, CP, Trios, Gene:IRF6



Explore Project Data

The samples for this study come from three primary ancestry groups: Europeans (from the United States, Denmark, Hungary, Spain, and Turkey), Asians (from China, the Philippines, and India), Central/South Americans (from Puerto Rico, Guatemala, Colombia, and Argentina), and Africans (from Nigeria and Ethiopia). The subjects have been assessed for various cleft types and related subclinical phenotypes. Our results database contains the GWAS results from this study of OFCs and can be explored as one study or subset into separate groups.

Custom Plots

Please select one option from each of the categories. You must select one option from each available category to be able to submit the request.

Genotype Data <input checked="" type="radio"/> Genotyped <input type="radio"/> Imputed	Phenotype <input checked="" type="radio"/> Cleft Palate (CP) <input type="radio"/> Cleft Lip With Or Without Cleft Palate (CL/P)
Ancestry Groups <input checked="" type="radio"/> All <input type="radio"/> Asian <input type="radio"/> European <input type="radio"/> Central/South Americans	Analysis Group <input checked="" type="radio"/> Trios <input type="radio"/> Case-Control <input type="radio"/> Meta-analysis
	Output <input type="radio"/> Manhattan Plot <input checked="" type="radio"/> Region/Locus <div style="border: 1px solid black; padding: 5px; margin-top: 10px;"><p>Please Select an option</p><p>Explore custom regions</p><p><input type="radio"/> Gene <input type="radio"/> SNP <input checked="" type="radio"/> Ch: BP-BP</p><p><input type="text" value="chr1:209458967-210479"/></p></div>

Submit

Reset



Explore Project Data

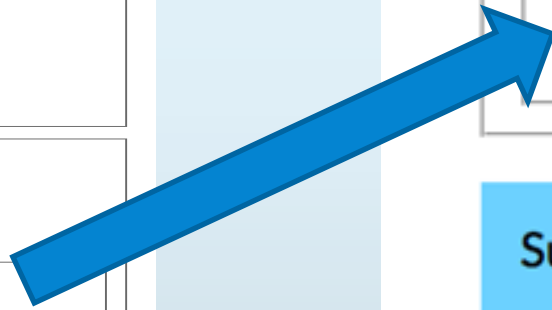
The samples for this study come from three primary ancestry groups: Europeans (from the United States, Denmark, Hungary, Spain, and Turkey), Asians (from China, the Philippines, and India), Central/South Americans (from Puerto Rico, Guatemala, Colombia, and Argentina), and Africans (from Nigeria and Ethiopia). The subjects have been assessed for various cleft types and related subclinical phenotypes. Our results database contains the GWAS results from this study of OFCs and can be explored as one study or subset into separate groups.

Custom Plots

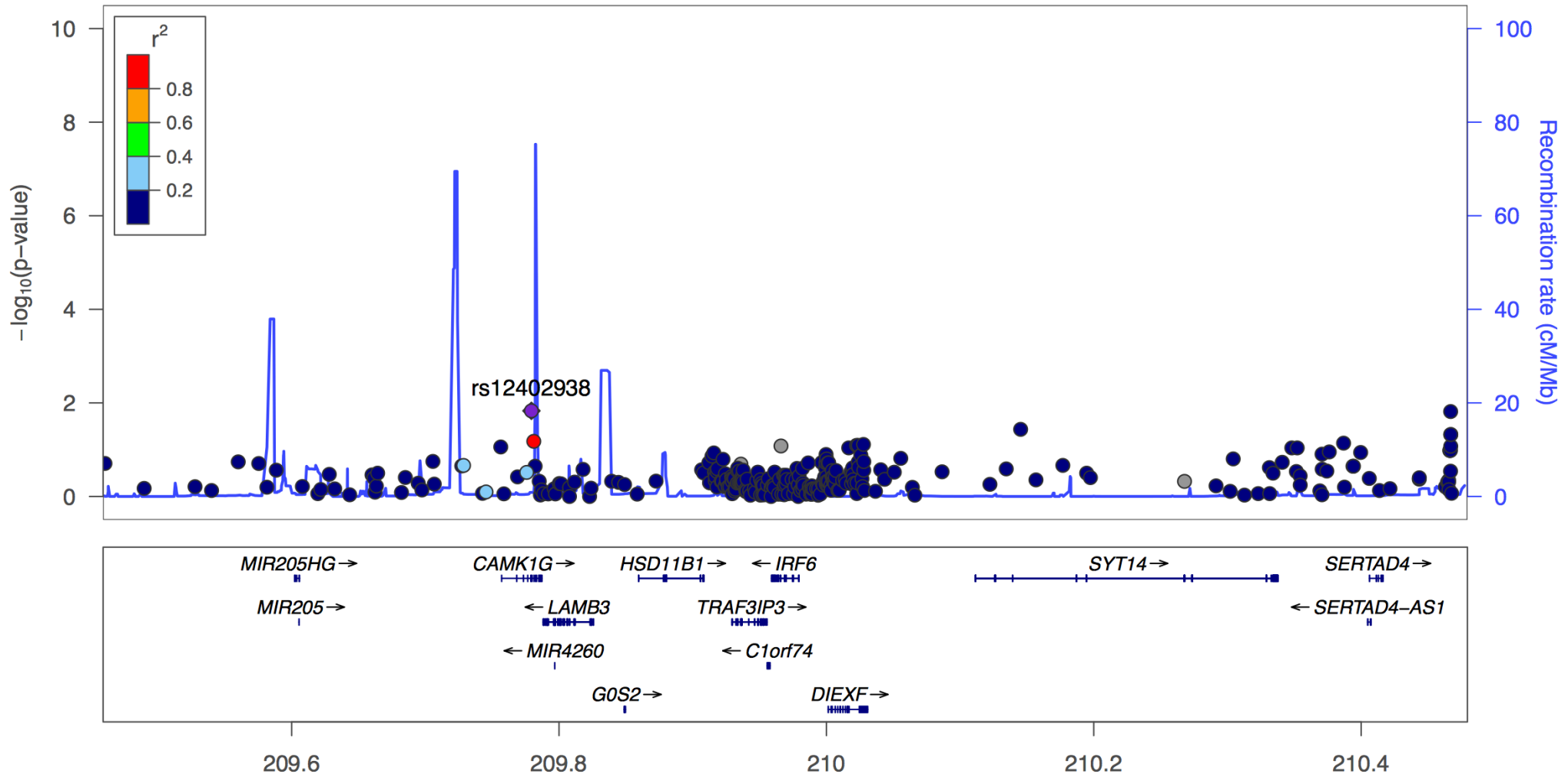
Please select one option from each of the categories. You must select one option from each available category to be able to submit the request.

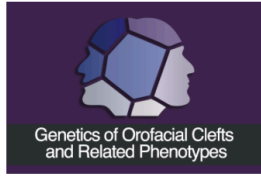
Genotype Data <input checked="" type="radio"/> Genotyped <input type="radio"/> Imputed	Phenotype <input checked="" type="radio"/> Cleft Palate (CP) <input type="radio"/> Cleft Lip With Or Without Cleft Palate (CL/P)
Ancestry Groups <input checked="" type="radio"/> All <input type="radio"/> Asian <input type="radio"/> European <input type="radio"/> Central/South Americans	Analysis Group <input checked="" type="radio"/> Trios <input type="radio"/> Case-Control <input type="radio"/> Meta-analysis
Output <input type="radio"/> Manhattan Plot <input checked="" type="radio"/> Region/Locus	
Please Select an option Explore custom regions <input type="radio"/> Gene <input type="radio"/> SNP <input checked="" type="radio"/> Ch: BP-BP <input type="text" value="chr1:209458967-210479"/>	
<input type="button" value="Submit"/> <input type="button" value="Reset"/>	

<input type="radio"/> Manhattan Plot <input checked="" type="radio"/> Region/Locus
Please Select an option Explore custom regions <input type="radio"/> Gene <input type="radio"/> SNP <input checked="" type="radio"/> Ch: BP-BP <input type="text" value="chr1:209458967-210479"/>
<input type="button" value="Submit"/> <input type="button" value="Reset"/>



Geno HG19, All Ancestry, CP, Trios, Chr Bp-Bp: chr1 209458967-210479520





Project Description

Methods

Descriptive Statistics

Explore Project Data

Publications

dbGaP Study Accession: [phs000774.v1.p1](#).

Principal Investigator: Mary L. Marazita, University of Pittsburgh, Pittsburgh, PA, USA

Funding Sources:

X01-HG00784 Genetics of Orofacial Clefts and Related Phenotypes. National Human Genome Research Institute, CIDR Genotyping, National Institutes of Health, Bethesda, MD, USA

R01-DE-016148, Extending the Phenotype of Nonsyndromic Orofacial Clefts. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R01-DE-012472, HOMOZYGOSITY MAPPING OF ORAL-FACIAL CLEFTS IN TURKEY. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R01-DE-009886, GENE MAPPING STUDIES OF ORAL/FACIAL CLEFTS - CHINA/INDIA. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

P50-DE-016215, CRANIOFACIAL ANOMALIES RESEARCH CENTER. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

K99-DE-022378, GENETIC STUDIES OF NONSYNDROMIC CLEFTS IN POPULATIONS OF AFRICAN DESCENT. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R37-DE-008559, MOLECULAR GENETIC EPIDEMIOLOGY OF CLEFT LIP AND PALATE. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R01-DE-014667, CLEFT LIP GENETICS: A MULTI CENTER INTERNATIONAL CONSORTIUM. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R01-DD-000295, HEALTH OUTCOMES AND IMPROVED PHENOTYPIC CHARACTERIZATION OF CLEFT LIP AND PALATE. National Center on Birth Defects and Developmental Disabilities, Centers for Disease Control and Prevention, Atlanta, GA, USA

R21-DE-016930, PLANNING INTERNATIONAL OROFACIAL CLEFT GENETIC STUDIES. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

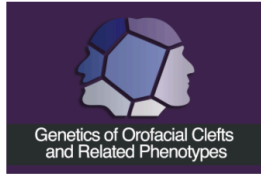
The current study is the culmination of many years of ongoing collaborations among the study investigators (for complete list of investigators, see [dbGaP](#)). Molecular genetic studies of orofacial clefts (OFC) by members of this research consortium began in the 1980's, followed by genome-wide linkage studies in the 1990's-early 2000's, and then genome-wide association studies in the 2010's. A rich phenotyping approach was first added to the consortium in 1999 as a project led by Dr. Mary Marazita under a center grant headed by Dr. Jeff Murray. Additional sites were added throughout the 2000's, until multiple populations and a large number of individuals (~12,000) comprise the current study population for genetic studies of OFCs and related subclinical phenotypes—subtle features believed to represent mild manifestations of the same underlying genetic susceptibility responsible for OFCs. As such, their inclusion in case-control and family-based genetic studies can help to clarify and refine the relationship between genotype and phenotype. Information on how to obtain individual level data is available on [dbGaP](#).



© 2014-2016 University of Pittsburgh.

This project is made possible by funding from NIDCR.
Grant #: U01-DE024425





Genetics of Orofacial Clefts and Related Phenotypes

Project Description Methods Descriptive Statistics Explore Project Data Publications

dbGaP Study Accession: [phs000774.v1.p1](#).

Principal Investigator: Mary L. Marazita, University of Pittsburgh, Pittsburgh, PA, USA
Funding Sources:

X01-HG00784 Genetics of Orofacial Clefts and Related Phenotypes. National Human Genome Research Institute, CIDR Genotyping, National Institutes of Health, Bethesda, MD, USA

R01-DE-016148, Extending the Phenotype of Nonsyndromic Orofacial Clefts. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R01-DE-012472, HOMOZYGOSITY MAPPING OF ORAL-FACIAL CLEFTS IN TURKEY. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R01-DE-009886, GENE MAPPING STUDIES OF ORAL/FACIAL CLEFTS - CHINA/INDIA. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

P50-DE-016215, CRANIOFACIAL ANOMALIES RESEARCH CENTER. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

K99-DE-022378, GENETIC STUDIES OF NONSYNDROMIC CLEFTS IN POPULATIONS OF AFRICAN DESCENT. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R37-DE-008559, MOLECULAR GENETIC EPIDEMIOLOGY OF CLEFT LIP AND PALATE. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

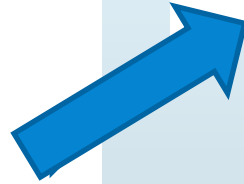
R01-DE-014667, CLEFT LIP GENETICS: A MULTI CENTER INTERNATIONAL CONSORTIUM. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R01-DD-000295, HEALTH OUTCOMES AND IMPROVED PHENOTYPIC CHARACTERIZATION OF CLEFT LIP AND PALATE. National Center on Birth Defects and Developmental Disabilities, Centers for Disease Control and Prevention, Atlanta, GA, USA

R21-DE-016930, PLANNING INTERNATIONAL OROFACIAL CLEFT GENETIC STUDIES. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

The current study is the culmination of many years of ongoing collaborations among the study investigators (for complete list of investigators, see [dbGaP](#)). Molecular genetic studies of orofacial clefts (OFC) by members of this research consortium began in the 1980's, followed by genome-wide linkage studies in the 1990's-early 2000's, and then genome-wide association studies in the 2010's. A rich phenotyping approach was first added to the consortium in 1999 as a project led by Dr. Mary Marazita under a center grant headed by Dr. Jeff Murray. Additional sites were added throughout the 2000's, until multiple populations and a large number of individuals (~12,000) comprise the current study population for genetic studies of OFCs and related subclinical phenotypes—subtle features believed to represent mild manifestations of the same underlying genetic susceptibility responsible for OFCs. As such, their inclusion in case-control and family-based genetic studies can help to clarify and refine the relationship between genotype and phenotype. Information on how to obtain individual level data is available on [dbGaP](#).

The current study is the culmination of many years of ongoing collaborations among the study investigators (for complete list of investigators, see [dbGaP](#)). Molecular genetic studies of orofacial clefts (OFC) by members of this research consortium began in the 1980's, followed by genome-wide linkage studies in the 1990's-early 2000's, and then genome-wide association studies in the 2010's. A rich phenotyping approach was first added to the consortium in 1999 as a project led by Dr. Mary Marazita under a center grant headed by Dr. Jeff Murray. Additional sites were added throughout the 2000's, until multiple populations and a large number of individuals (~12,000) comprise the current study population for genetic studies of OFCs and related subclinical phenotypes—subtle features believed to represent mild manifestations of the same underlying genetic susceptibility responsible for OFCs. As such, their inclusion in case-control and family-based genetic studies can help to clarify and refine the relationship between genotype and phenotype. Information on how to obtain individual level data is available on [dbGaP](#).



© 2014-2016 University of Pittsburgh.

This project is made possible by funding from NIDCR.
Grant #: U01-DE024425





Center for Craniofacial and Dental Genetics (CCDG): Genetics of Orofacial Clefts and Related Phenotypes

dbGaP Study Accession: phs000774.v1.p1

Show BioProject list

Study | Variables | Documents | Analyses | Datasets | Molecular Data

Jump to: [Authorized Access](#) | [Attribution](#) | [Authorized Requests](#)

Study Description

The purpose of this study is to investigate the genetics of orofacial clefts (OFCs) in a large study population, and importantly, to incorporate subclinical phenotypic features into these studies.

Orofacial clefts (OFCs) comprise a significant fraction of human birth defects (about 1/700 live births (Rahimov et al. 2012) and represent a major public health challenge, as individuals with these anomalies require surgical, nutritional, dental, speech, medical and behavioral interventions, thus imposing a substantial economic and personal burden (Berk and Marazita 2002*). The most common forms include OFCs of the lip alone (CL, Figure 1A), CL plus cleft palate (CL+CP, Figure 1B) or of the palate only (CP, Figure 1C). Individuals born with OFC may have their first surgical repair at age 3 months, but this initial surgery is just the beginning of a lifetime of health burdens. An individual born with an OFC has a hospital use rate increased for most ages (up to 233% increase for children ages 0-10 years and 16% for middle aged adults (Wehby et al. 2012). Healthcare costs for children with OFCs are estimated to be 800% greater compared with their unaffected peers (Boulet et al. 2009). Data from Denmark show that people born with CL with or without CP (CL/P) have an increased mortality up to age 55, which may be attributed to an increased risk of suicide and/or certain cancers (Christensen et al. 2004). The focus of most OFC genetic research has been CL and/or CP. Furthermore, the majority of OFC, i.e. about 70% of CL/P and 50% of CP is considered "nonsyndromic" (Jones 1988), i.e. isolated anomalies with no other apparent cognitive or structural abnormalities.

Figure 1 Sample OFC Types



A: Bilateral Cleft Lip; B=Cleft Lip plus Cleft Palate; C:Cleft Palate Alone

The factors leading to the majority of nonsyndromic OFCs are still unclear, particularly at an individual family level. As is true for many complex traits, substantial progress in gene identification has occurred in the OFC field in the last two years (Dixon et al. 2011; Marazita 2012). Genome Wide Association Studies (GWAS) and sequencing studies to date by our research team and others have focused on genetic risk factors for overt CL/P and CPO-and have been very successful. A major finding from this work is that OFCs exhibit significant genetic heterogeneity, i.e., multiple genetic regions have been implicated (Beaty et al. 2010; Ludwig et al. 2012). Thus, approaches are needed to understand this genetic heterogeneity. Are there GxG interactions at work? Are there subsets of families, each due to a different gene? Our research group has shown that a promising approach to dissect the etiology of OFC is to focus on subclinical phenotypic features within entire cleft families (not just in affected cases, but also in their non-cleft relatives). These subtle features are believed to represent mild manifestations of the same underlying genetic susceptibility responsible for OFCs; as such, their inclusion in case-control and family-based genetic studies can help to clarify and refine the relationship between genotype and phenotype.

The study population comprises a large number of families and individuals (~12,000 individuals) from multiple populations worldwide (Caucasians from the US and Europe, Asians from China and the Philippines, Mixed Native American/Caucasians from South America, and Africans from Nigeria and Ethiopia). There are cases, case families (nuclear families and extended kindreds), as well as controls with no history of OFC nor other developmental defects.

*Berk NW, Marazita ML (2002) Costs of Cleft Lip and Palate: Personal and Societal Implications. In: Wyszynski DF (ed) Cleft Lip and Palate: From Origin to Treatment. Oxford University Press, Inc., New York, pp 458-467.

- Study Types: Parent-Offspring, Nuclear Families, Extended Pedigrees
- Number of study subjects that have individual level data available through Authorized Access: 11925
 - 11925 phenotyped subjects

Important Links and Information

- Request access via [Authorized Access](#)
 - [Instructions](#) for requestors
 - [Data Use Certification \(DUC\) Agreement](#)
- [Talking Glossary of Genetic Terms](#)

Authorized Access

- **Data access provided by:** [dbGaP Authorized Access](#)
- **Release Date:** November 02, 2015
- **Embargo Release Date:** November 02, 2015
- [Data Use Certification Requirements \(DUC\)](#)
- **Use Restrictions**

Consent group	Is IRB required?	Data Access Committee	Number of participants
Disease-Specific (Craniofacial Research) 0	No	Joint NIAMS-NIDCR Data Access Committee (jardedac@mail.nih.gov)	11925

- [List of components](#) downloadable from [Authorized Access](#)

Publicly Available Data (Public ftp)

Connect to the [public download site](#). The site contains release notes and manifests. If available, the site also contains data dictionaries, variable summaries, documents, and truncated analyses.

Study Inclusion/Exclusion Criteria

Inclusion criteria for Families with OFCs:

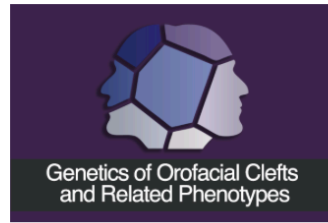
- Case diagnosed with nonsyndromic OFC (CL/P or CPO).
- At least 2 additional family members' participation.
- Male or female, aged 1 month to 110 years.
- Provide signed and dated informed consent form.
- Willing to comply with all study procedures and be available for the duration of the study.

Inclusion criteria for Controls:

- Male or female, aged 1 month to 110 years.
- No known personal or family history of OFCs, genetic conditions, or birth defects.
- No personal history of facial surgery.
- In good general health.
- Provide signed and dated informed consent form.
- Willing to comply with all study procedures and be available for the duration of the study.

Exclusion criteria - individuals who met any of the following criteria were excluded from participation in this study:

- An affected case with a genetic syndrome or multiple congenital anomalies.
- An affected case without participation of other family members.
- Persons who were adopted and cannot report family history.
- For controls, a positive history of OFC, genetic syndrome/defects in themselves or family.
- For controls, a previous facial surgery.
- Male controls may be excluded from certain phenotypes if it is determined that their facial hair hinders the ability to collect accurate facial measurements.



- Project Description
- Methods
- Descriptive Statistics
- Explore Project Data
- Publications

dbGaP Study Accession: [phs000774.v1.p1](#)

Principal Investigator: Mary L. Marazita, University of Pittsburgh, Pittsburgh, PA, USA

Funding Sources:

X01-HG00784 Genetics of Orofacial Clefts and Related Phenotypes. National Human Genome Research Institute, CIDR Genotyping, National Institutes of Health, Bethesda, MD, USA

R01-DE-016148, Extending the Phenotype of Nonsyndromic Orofacial Clefts. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R01-DE-012472, HOMOZYGOSITY MAPPING OF ORAL-FACIAL CLEFTS IN TURKEY. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R01-DE-009886, GENE MAPPING STUDIES OF ORAL/FACIAL CLEFTS - CHINA/INDIA. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

P50-DE-016215, CRANIOFACIAL ANOMALIES RESEARCH CENTER. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

K99-DE-022378, GENETIC STUDIES OF NONSYNDROMIC CLEFTS IN POPULATIONS OF AFRICAN DESCENT. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R37-DE-008559, MOLECULAR GENETIC EPIDEMIOLOGY OF CLEFT LIP AND PALATE. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R01-DE-014667, CLEFT LIP GENETICS: A MULTI CENTER INTERNATIONAL CONSORTIUM. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

R01-DD-000295, HEALTH OUTCOMES AND IMPROVED PHENOTYPIC CHARACTERIZATION OF CLEFT LIP AND PALATE. National Center on Birth Defects and Developmental Disabilities, Centers for Disease Control and Prevention, Atlanta, GA, USA

R21-DE-016930, PLANNING INTERNATIONAL OROFACIAL CLEFT GENETIC STUDIES. National Institute of Dental and Craniofacial Research, National Institutes of Health, Bethesda, MD, USA

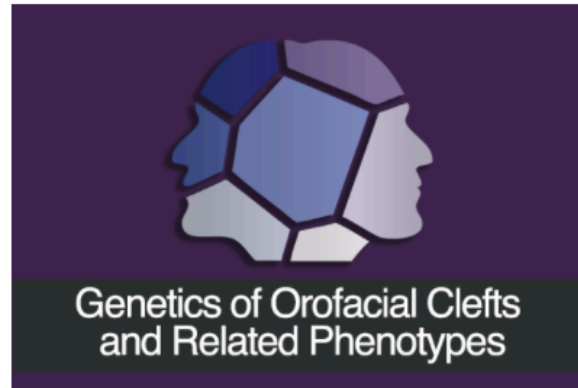
The current study is the culmination of many years of ongoing collaborations among the study investigators (for complete list of investigators, see [dbGaP](#)). Molecular genetic studies of orofacial clefts (OFC) by members of this research consortium began in the 1980's, followed by genome-wide linkage studies in the 1990's-early 2000's, and then genome-wide association studies in the 2010's. A rich phenotyping approach was first added to the consortium in 1999 as a project led by Dr. Mary Marazita under a center grant headed by Dr. Jeff Murray. Additional sites were added throughout the 2000's, until multiple populations and a large number of individuals (~12,000) comprise the current study population for genetic studies of OFCs and related subclinical phenotypes—subtle features believed to represent mild manifestations of the same underlying genetic susceptibility responsible for OFCs. As such, their inclusion in case-control and family-based genetic studies can help to clarify and refine the relationship between genotype and phenotype. Information on how to obtain individual level data is available on [dbGaP](#).



© 2014-2016 University of Pittsburgh.

This project is made possible by funding from NIDCR.
Grant #: U01-DE024425





Project
Description

Methods

Descriptive
Statistics

Explore
Project Data

Publications

Publications

[PubMed list](#) for all publications associated with this project.



© 2014-2016 University of Pittsburgh.

This project is made possible by funding from NIDCR.
Grant #: U01-DE024425



Abstract

Send to:

Am J Hum Genet. 2016 Apr 7;98(4):744-54. doi: 10.1016/j.ajhg.2016.02.014. Epub 2016 Mar 24.

A Genome-wide Association Study of Nonsyndromic Cleft Palate Identifies an Etiologic Missense Variant in GRHL3.

Leslie EJ¹, Liu H², Carlson JC³, Shaffer JR⁴, Feingold E⁵, Wehby G⁶, Laurie CA⁷, Jain D⁷, Laurie CC⁷, Doheny KF⁸, McHenry T¹, Resick J¹, Sanchez C¹, Jacobs J¹, Emanuele B¹, Vieira AR⁴, Neiswanger K¹, Standley J⁹, Czeizel AE¹⁰, Deleviannis F¹¹, Christensen K¹², Munger RG¹³, Lie RT¹⁴, Wilcox A¹⁵, Romitti PA¹⁶, Field LL¹⁷, Padilla CD¹⁸, Cutionco-de la Paz EM¹⁹, Lidral AC²⁰, Valencia-Ramirez LC²¹, Lopez-Palacio AM²², Valencia DR²³, Arcos-Burgos M²⁴, Castilla EE²⁵, Mereb JC²⁶, Poletta FA²⁵, Orioli IM²⁷, Carvalho FM²⁸, Hecht JT²⁹, Blanton SH³⁰, Buxó CJ³¹, Butali A³², Mossey PA³³, Adeyemo WL³⁴, James O³⁴, Braimah RO³⁵, Aregbesola BS³⁵, Eshete MA³⁶, Deribew M³⁶, Koruyucu M³⁷, Seymen F³⁷, Ma L³⁸, de Salamanca JE³⁹, Weinberg SM¹, Moreno L²⁰, Corneli RA⁴⁰, Murray JC⁹, Marazita ML⁴¹.

Author information

Abstract

Cleft palate (CP) is a common birth defect occurring in 1 in 2,500 live births. Approximately half of infants with CP have a syndromic form, exhibiting other physical and cognitive disabilities. The other half have nonsyndromic CP, and to date, few genes associated with risk for nonsyndromic CP have been characterized. To identify such risk factors, we performed a genome-wide association study of this disorder. We discovered a genome-wide significant association with a missense variant in GRHL3 (p.Thr454Met [c.1361C>T]; rs41268753; p = 4.08 × 10⁽⁻⁹⁾) and replicated the result in an independent sample of case and control subjects. In both the discovery and replication samples, rs41268753 conferred increased risk for CP (OR = 8.3, 95% CI 4.1-16.8; OR = 2.16, 95% CI 1.43-3.27, respectively). In luciferase transactivation assays, p.Thr454Met had about one-third of the activity of wild-type GRHL3, and in zebrafish embryos, perturbed periderm development. We conclude that this mutation is an etiologic variant for nonsyndromic CP and is one of few functional variants identified to date for nonsyndromic orofacial clefting. This finding advances our understanding of the genetic basis of craniofacial development and might ultimately lead to improvements in recurrence risk prediction, treatment, and prognosis.

Copyright © 2016 The American Society of Human Genetics. Published by Elsevier Inc. All rights reserved.

PMID: 27018472 [PubMed - in process] PMID: PMC4833215 [Available on 2016-10-07]



Grant Support

LinkOut - more resources

PubMed Commons

PubMed Commons home

0 comments

How to join PubMed Commons

Full text links



Save items

Add to Favorites

Similar articles

Sequencing the GRHL3 Coding Region Reveals Rare Truncating Mutati [Am J Hum Genet. 2016]

Review Toward an orofacial gene regulatory network. [Dev Dyn. 2016]

Rare functional variants in genome-wide association identified [Am J Med Genet A. 2014]

Analysis of susceptibility loci for nonsyndromic orofacial clefting in a [Am J Med Genet A. 2013]

Review Genetics of nonsyndromic orofacial clefts. [Cleft Palate Craniofac J. 2012]

See reviews...

See all...

Related information

Articles frequently viewed together

MedGen

References for this PMC Article

Recent Activity

Turn Off Clear

A Genome-wide Association Study of Nonsyndromic Cleft Palate Identifies PubMed

U01-HG005925 (0) PMC

U01-HG005925[All Fields] (0) PMC

See more...

You are here: NCBI > Literature > PubMed

Write to the Help Desk

GETTING STARTED

- NCBI Education
- NCBI Help Manual
- NCBI Handbook
- Training & Tutorials
- Submit Data

RESOURCES

- Chemicals & Bioassays
- Data & Software
- DNA & RNA
- Domains & Structures
- Genes & Expression
- Genetics & Medicine
- Genomes & Maps

POPULAR

- PubMed
- Bookshelf
- PubMed Central
- PubMed Health
- BLAST
- Nucleotide
- Genome

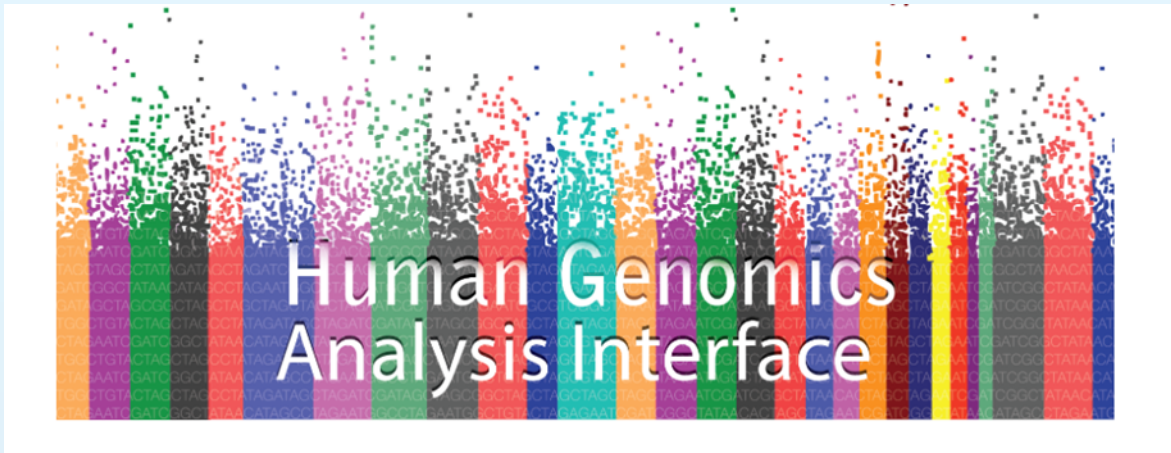
FEATURED

- Genetic Testing Registry
- PubMed Health
- GenBank
- Reference Sequences
- Gene Expression Omnibus
- Map Viewer
- Human Genome

NCBI INFORMATION

- About NCBI
- Research at NCBI
- NCBI News
- NCBI FTP Site
- NCBI on Facebook
- NCBI on Twitter
- NCBI on YouTube

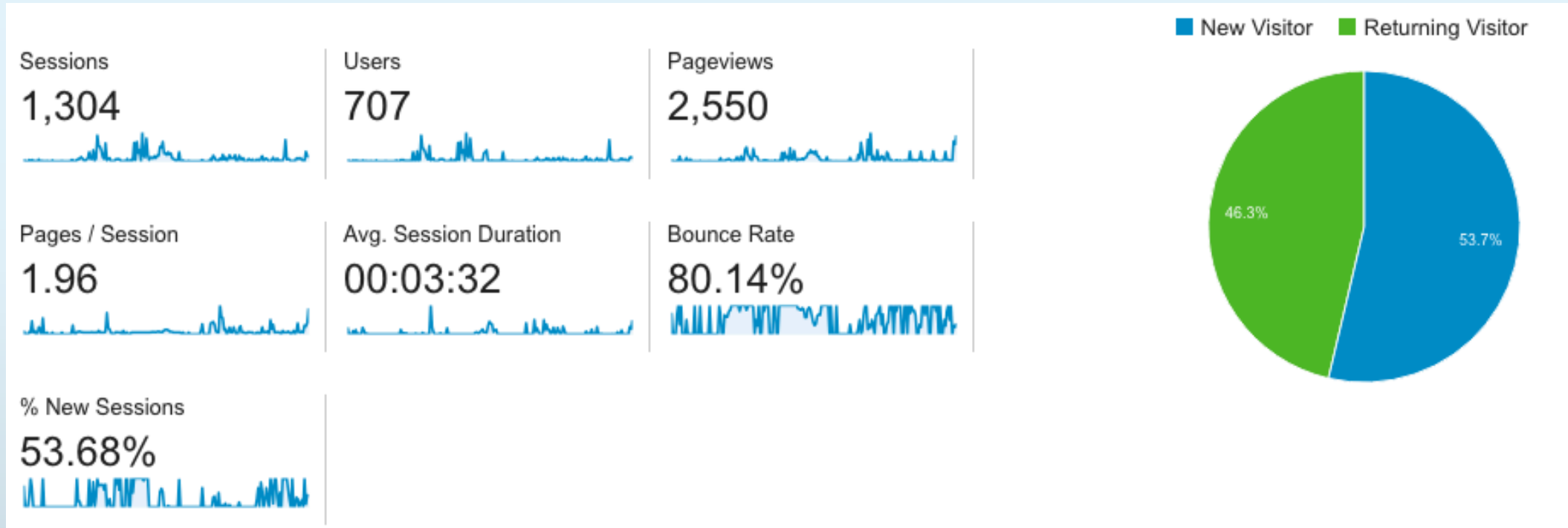























VISITS TO THE SITE



GOOGLE ANALYTICS (since October 2015)



GOOGLE ANALYTICS (since October 2015)

Country	Sessions	% Sessions
1.  United States	525	 40.26%
2.  Russia	279	 21.40%
3. (not set)	195	 14.95%
4.  China	41	 3.14%
5.  United Kingdom	30	 2.30%
6.  Japan	21	 1.61%
7.  Netherlands	21	 1.61%
8.  Germany	15	 1.15%
9.  South Korea	14	 1.07%
10.  Spain	11	 0.84%

FUTURE PLANS

- Complete existing projects
 - Other potential OFC GWAS: CHoP, Chinese, others
 - US Facial Variation, Dental
- Seek additional projects
 - UK Facial Variation
 - Other oral/dental
- Incorporate additional functionality
- Citing tool in papers
- Possible dbGaP link to tool



FUTURE PLANS: FaceBase HUB

- Add links from animal model pages to this human page and vice versa
 - E.g. for “look-up” of gene or region of interest
- FaceBase human data projects: any aspects that could be incorporated into this tool?
- Suggestions, requests?



ACKNOWLEDGEMENTS

- Funding NIDCR

- U01-DE024425

- Faculty:

Elizabeth Leslie

Eleanor Feingold

Harry Hochheiser

- Staff/Students:

Annette Werner

Jenna Carlson

Myoungkeun Lee

Lance Kennelty

Tom Maher

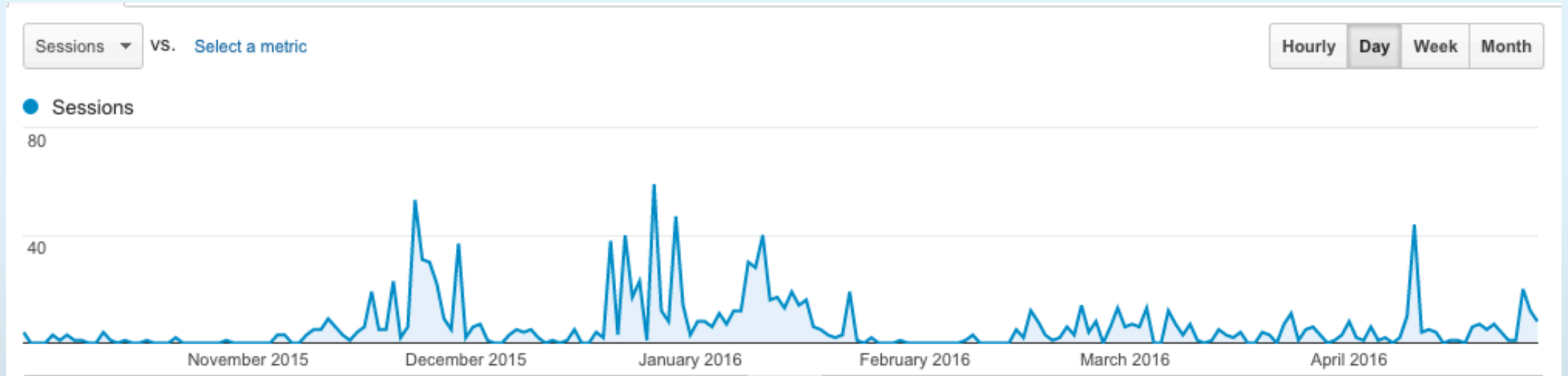
Jim Gallagher

Justin Stickel





GOOGLE ANALYTICS (since October 2015)



FaceBase2 Production cluster At University of Pittsburgh's Network Operations Center

