Introduction to Bioinformatics

Richard H. Scheuermann, Ph.D. Director of Informatics JCVI







Outline

- What is Bioinformatics?
 - Some definitions
 - Data types and analysis objectives
- Big Data
 - The Big Data value proposition
- The Power of Bioinformatics
 - Extracting knowledge from data
 - DMID Systems Biology data in the Bioinformatics Resource Centers







What is Bioinformatics?

- And related terms biomedical informatics, computational biology, systems biology
- Wikipedia
 - Bioinformatics: an interdisciplinary field that develops and improves on methods for storing, retrieving, organizing and analyzing biological data. A major activity in bioinformatics is to develop software tools to generate useful biological knowledge.
- NIH Biomedical Information Science and Technology Initiative Consortium (BISTIC)
 - Bioinformatics: Research, development, or application of computational tools and approaches for expanding the use of biological, medical, behavioral or health data, including those to acquire, store, organize, archive, analyze, or visualize such data.
 - Computational Biology: The development and application of data-analytical and theoretical methods, mathematical modeling and computational simulation techniques to the study of biological, behavioral, and social systems.







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Biological data types and analysis objectives

Genomics

- Nucleotide genome sequences, metagenomic sequences
- Gene finding, functional annotation, sequence alignment, homology determination, comparative analysis, phylogenetic inferencing, association analysis, mutation functional prediction, species distribution analysis

Transcriptomics

- RNA expression levels, transcription factor binding, chromatin structure information
- Differential expression, clustering, functional enrichment, transcriptional regulation/causal reasoning

Proteomics

- Proteins levels, protein structures, protein interactions
- Protein identification, protein functional predictions, structural predictions, structural comparison, molecular dynamic simulation, mutation functional prediction, docking predictions, network analysis

Metabolomics

- Metabolite/small molecule levels
- Pathway/network analysis

Imaging

- Microscopy images, MRI images, CT scans
- Feature extraction, high content screening

• Cytometry

- Cell levels, cell phenotypes
- Cell population clustering, cell biomarker discovery

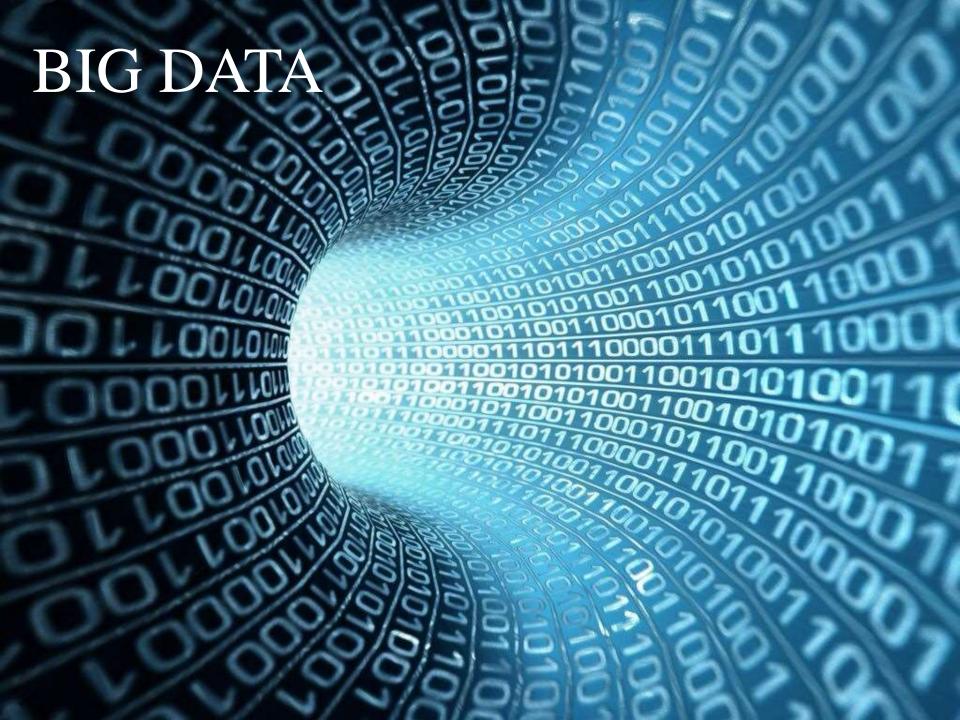
Systems biology

- All of the above
- Network analysis, causal reasoning, reverse causal reasoning, drug target prediction, regulatory network analysis, information flow, population dynamics, modeling and simulation













FUNDING OPPORTUNITIES & NOTICES

WORKSHOPS

NEWS

ABOUT BD2K

FAQs



The NIH Big Data to Knowledge (BD2K) announces funding opportunity for

CENTERS OF EXCELLENCE FOR BIG DATA COMPUTING IN THE BIOMEDICAL SCIENCES

LEARN MORE

The mission of the NIH Big Data to Knowledge (BD2K) initiative is to enable biomedical scientists to capitalize more fully on the Big Data being generated by those research communities. With advances in technologies, these investigators are increasingly generating and using large, complex, and diverse datasets. Consequently, the biomedical research enterprise is increasingly becoming data-intensive and data-driven. However, the ability of researchers to locate, analyze, and use Big Data (and more generally all biomedical and behavioral data) is often limited for reasons related to access to relevant software and tools, expertise, and other factors. BD2K aims to develop the new approaches, standards, methods, tools, software, and competencies that will enhance the use of biomedical Big Data by supporting research, implementation, and training in data science and other relevant fields that will lead to:

Read more

WORKSHOPS



Frameworks for Community-Based Standards Efforts

September 25 - 26, 2013

More Workshops >

NEWS HIGHLIGHT

- NIH Names Dr. Philip E. Bourne First Associate Director for Data Science
 December 9, 2013
- NIH commits \$24 million annually for Big Data Centers of Excellence
 July 22, 2013
- NIH to recruit Associate Director for Data Science January 10, 2013
- NIH proposes critical initiatives to sustain future of U.S. biomedical research
 December 7, 2012

More News >

FUNDING OPPORTUNITIES & NOTICES

- Development of an NiH BD2K Data Discovery Index Coordination Consortium(U24)
 December 17, 2013
- BD2K-LINCS-Perturbation Data Coordination and Integration Center (DCIC) (U54)
 December 4, 2013
- Input on Development of Analysis Methods and Software for Big Data (RFI) August 8, 2013
- Centers of Excellence for Big Data Computing in the Biomedical Sciences (U54)
 July 22, 2013

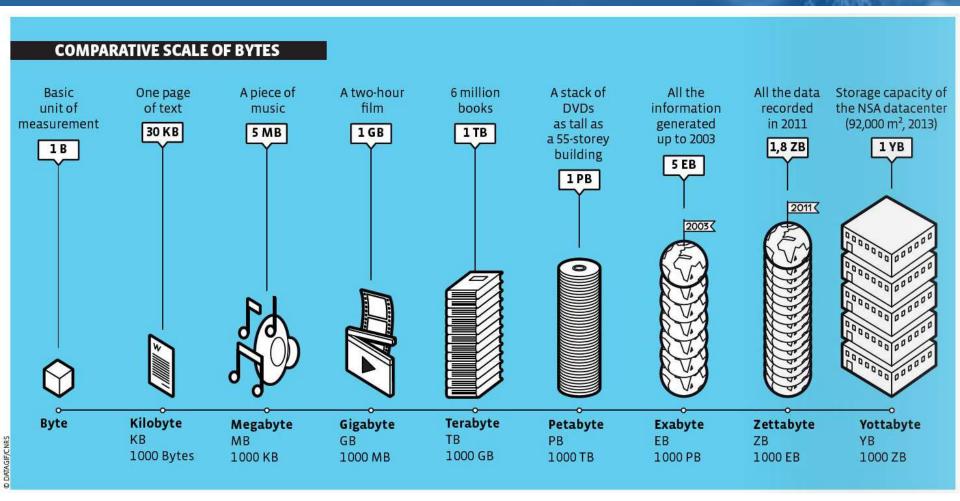
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Big Data Volumes

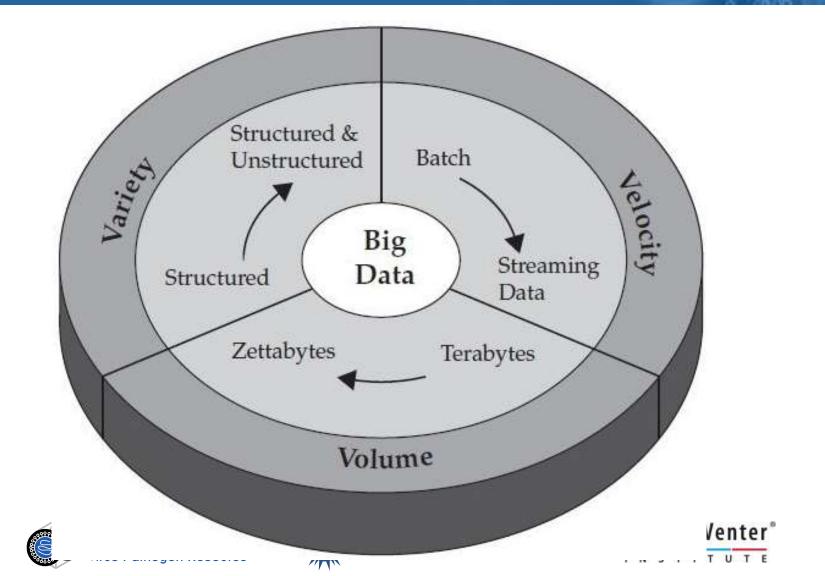




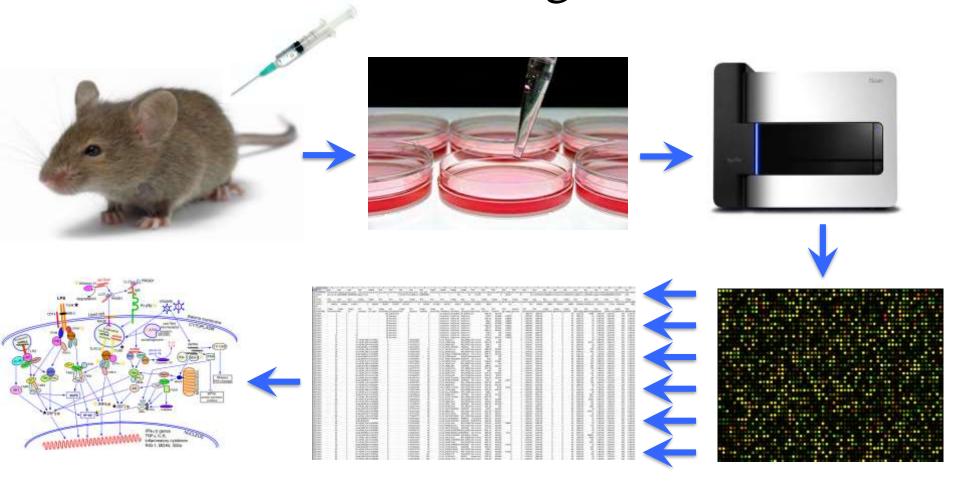




Big Data 3 V's



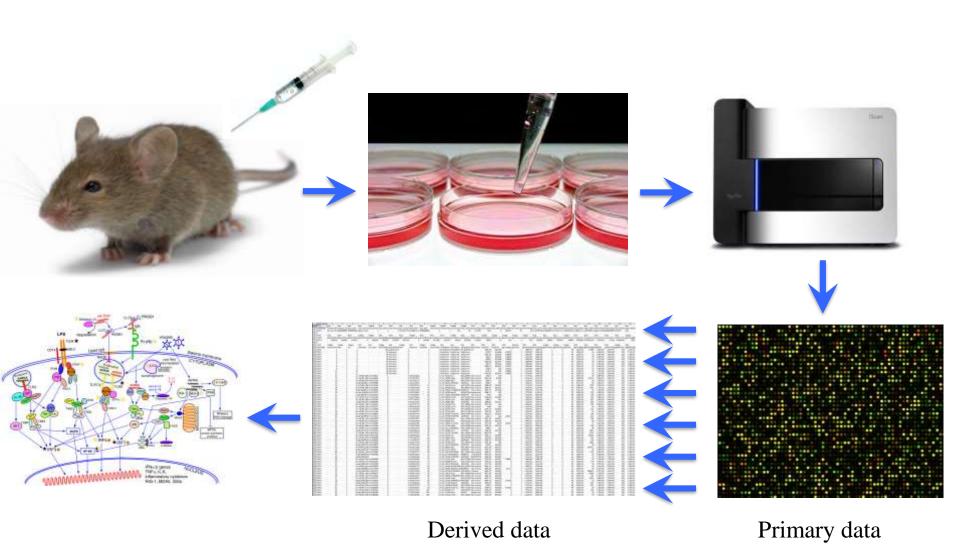
Data Levels in Biological Research









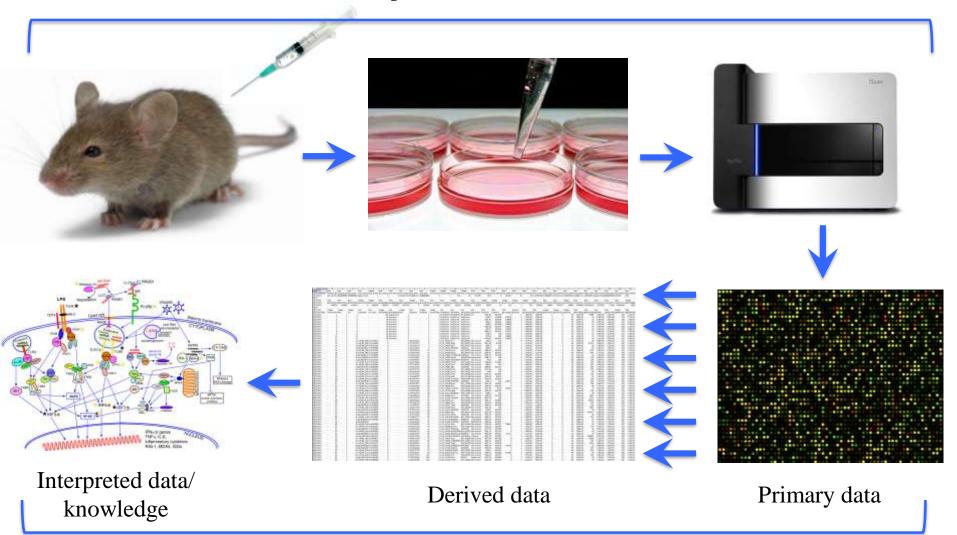








Experimental metadata



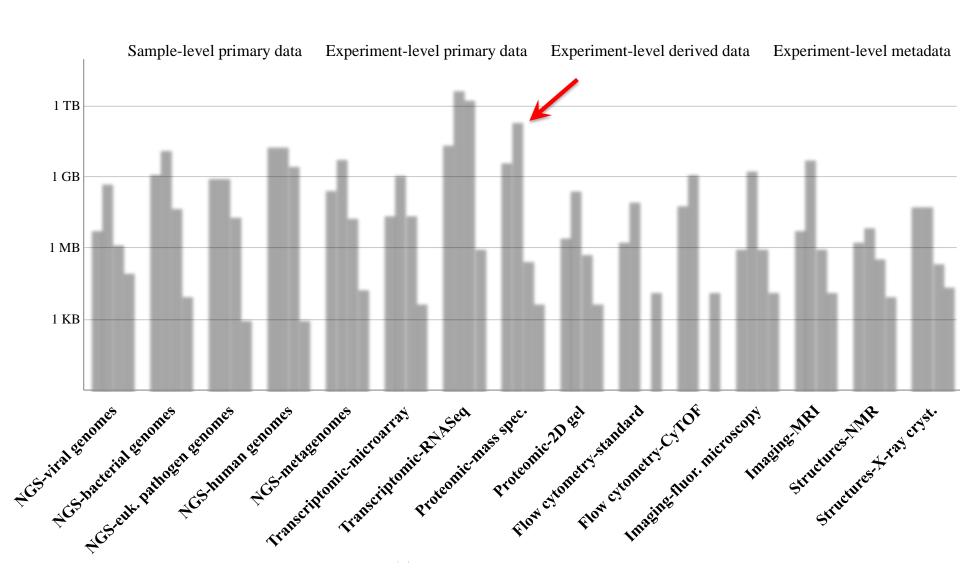
Analytical metadata



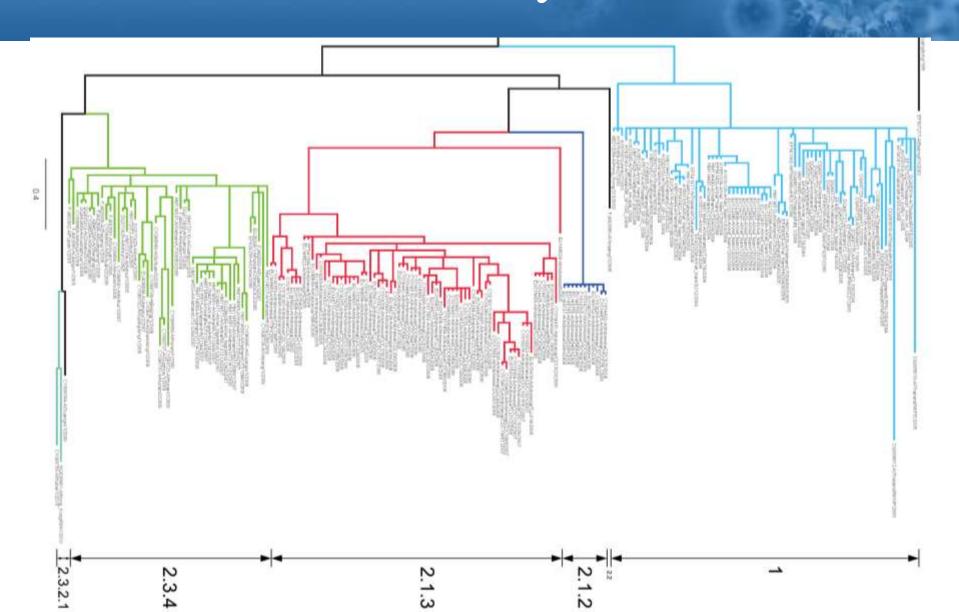




Big Data in Biology



Variety



No Variety

A/Boston/26/2008

A/Canada-MB/RV1975/2009 A/Michigan/30/2009 A/Lyon/969/2009 A/Mexico/InDRE13495/2009 A/Canada-SK/RV1767/2009 A/Canada-MB/RV2018/2009

A/England/348/2009 A/England/328/2009 A/Taiwan/90262/2011 A/Kanagawa/140/2009

A/England/345/2009 A/England/377/2009

A/England/349/2009 A/England/360/2009 A/England/364/2009

A/England/342/2009 A/England/374/2009

A/England/399/2009

A/England/350/2009

A/California/07/2009 A/California/08/2009 A/Beijing/16/2009

A/Helsinki/490/2013

A/Helsinki/753/2013 A/Helsinki/979/2013

A/California/07/2009

A/California/07/2009

A/California/07/2009

A/California/07/2009

A/California/08/2009 A/England/201/2009

A/Hangzhou/04/2009 A/California/08/2009 A/California/04/2009

A/California/04/2009 A/Pennsylvania/09/2009

A/California/04/2009 A/Hangzhou/06/2009

A/Fukuoka-C/2/2009 A/Fukuoka-C/3/2009 A/Hangzhou/10/2009

A/Fukuoka-C/1/2009

A/California/04/2009 A/Kagoshima/1/2009

Big Data

Volume + Variety = Value

Variety = Metadata







* NIAID/DMID Genomics Program

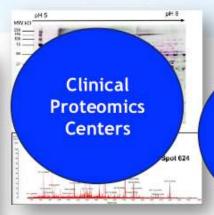
Sequencing

Genomic Sequencing Centers

Functional Genomics

Functional Genomic Research Centers Fooder and Control of Control

Proteomics



Structural Genomics

Structural Genomics Centers Systems Biology

Systems Biology Centers

Bioinformatics Resource Centers

Genomic Research Resources

Genomic/Omics Data Sets, Databases, Bioinformatics Tools, Biomarkers, 3D Structures, Protein Clones, Predictive Models

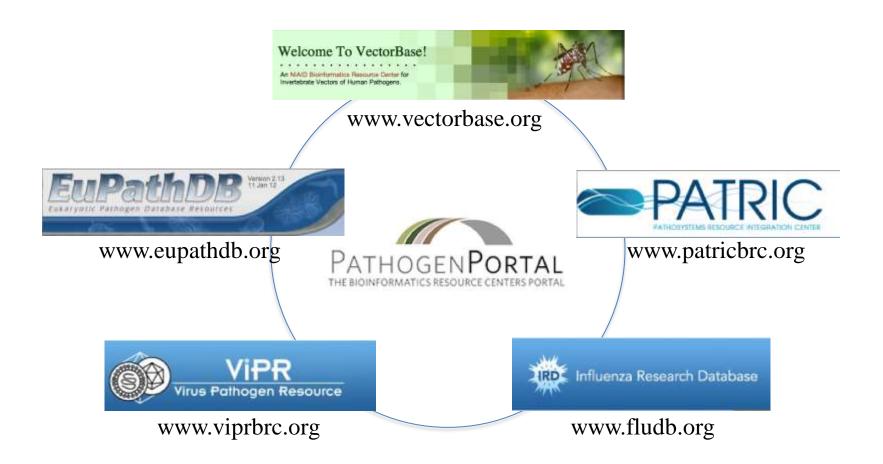


To address key questions in microbiology and infectious disease



To identify new targets and develop new strategies for vaccines, diagnostics and therapeutics

Bioinformatics Resource Centers (BRCs)











NIAID Home

Health & Research A to Z

Labs & Scientific Resources

Funding

About NIAID

News & Events

NIAID > Labs & Scientific Resources > Resources for Researchers > DMID Resources for Researchers > Systems Biology

Resources for Researchers

DMID Resources for Researchers

Systems Biology

News Releases

Related Resources

Systems Biology Working Group

Data Sharing Guidelines

Systems Biology for Infectious Diseases Research

What services do the programs provide?

The NIAID program in Systems Biology for Infectious Diseases Research utilizes a combination of computational and experimental methodologies to conduct research projects to analyze, identify, quantify, model, and predict the overall dynamics of the network of cellular molecular components of I Order publications microbial organisms and their interactions with the host cells. The knowledge generated from the research projects, including research data, analytical software tools, computational models, experimental protocols, and reagents, is widely disseminated to the scientific community through publicly accessible databases and reagent repositories. The research findings will provide a deeper understanding of the overall complexity of the biological, biochemical and biophysical molecular processes in microbial organisms as well as how the molecular events within the pathogen lead to the initiation and progression of infectious disease.

Where are services provided?

The research activities are carried out by

- Battelle—Pacific Northwest National Laboratory (Systems Biology for EnteroPathogens)
- Institute for Systems Biology (Systems Influenza)
- Stanford University and the Broad Institute (TB Systems Biology)
- University of Washington (Systems Virology)

Access

Research and associated data, protocols, and computational and statistical models will be made freely and publicly available to the scientific community through the research centers websites within 4 weeks of publication, or within 1 year of generation, whichever comes first and as agreed upon by the Project Officer.

Reagents will be made available through the BEI Resources Repository (BEI) ...

Related Resources and Information

- Data Sharing Guiding Principles
- News and Announcements
- Related NIAID-Supported Services

Website Tools

Email this page

Print this page

Get email updates

Get plug-ins and viewers

Bookmark & share

Stay Connected









Social media privacy policy and disclaimers.

Contact Info

Valentina Di Francesco E-mail:

vdifrancesco@niaid.nih.gov

Highlight

Funding Opportunity: "OMICS" Technologies For Predictive Modeling of Infectious Diseases (U19)

Building on a Decade of Accomplishments: Report of the 2010 Blue Ribbon Panel on Genomics (PDF)

A Systems Biology Approach to Infectious Disease Research: Innovating the Pathogen-Host Research Paradigm[®], Feb 2011

New class of biomolecules triggered in response to respiratory virus infection. October 25, 2010

What is systems biology?[™] Feb. 2010

Systems Biology of Viral Infection

- Systems Virology (Michael Katze group, Univ. Washington)
 - Influenza H1N1 and H5N1 and SARS Coronavirus
 - Statistical models, algorithms and software, raw and processed gene expression data, and proteomics data
- Systems Influenza (Alan Aderem group, Institute for Systems Biology/Seattle Biomed)
 - Various influenza viruses
 - Microarray, mass spectrometry, and lipidomics data







Data Dissemination Working Group

Representatives from SysBio programs and

relevant BRCs

Jeremy Zucker

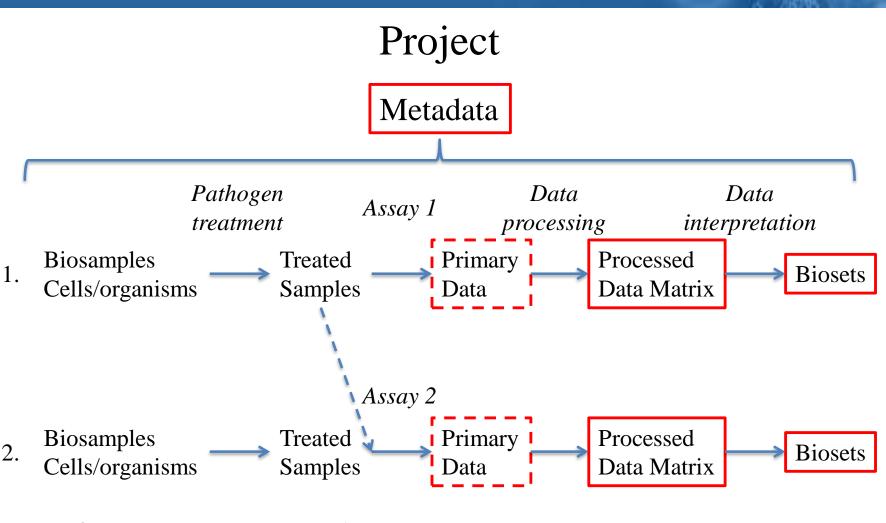








"Omics" Data Management









Strategy for Handling "Omics" Data

- "Omics" data management (MIBBI)
 - Project metadata (1 template)
 - Title, PI, abstract, publications
 - Experiment metadata (~6 templates)
 - Biosamples, treatments, reagents, protocols, subjects
 - Primary results data
 - Raw expression values
 - Processed data
 - Data matrix of fold changes and p-values
 - Data processing metadata (1 template)
 - Normalization and summarization methods
 - Interpreted results (Host factor biosets)
 - Interesting gene, protein and metabolite lists
 - Data interpretation metadata (1 template)
 - Fold change and p-value cutoffs used
- Visualize biosets in context of biological pathways and networks
- Statistical analysis of pathway/sub-network overrepresentation







Data Submission Workflows

Free text metadata GEO/PRIDE/PNNL/SRA/MetaboLights Primary results submission Study metadata pointer submission Experiment metadata ViPR/IRD/PATRIC Analysis metadata Processed data matrix pointer Host factor bioset Systems Biology sites









www.fludb.org

You are logged in as rscheuermann@jcvi.org

Sign Out

SEARCH DATA

ANALYZE & VISUALIZE

WORKBENCH

SUBMIT DATA

Search

Search our comprehensive database for:

- Influenza segment and protein sequences
- Avian and non-human mammalian surveillance data
- Virus phenotypic characteristics
- Host Factor Data (Prototype)
- Immune epitope data
- 3D protein structures

Browse All Search Types

Analyze

Analyze data online:

- Align sequences
- Identify similar sequences (BLAST)
- Identify short peptides in flu proteins
- Identify point mutations in flu proteins
- Analyze Sequence Variation (SNP)
- Generate a phylogenetic tree

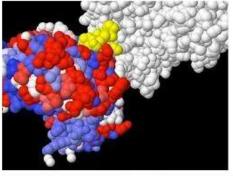
Browse All Tools

Save to Workbench

Use your workbench to:

- Store sequences or other data in working sets for future analysis
- Combine working sets
- Integrate IRD data with your laboratory data
- Store analysis results
- Share results

Highlights



St. Jude Children's Research Hospital CFIRS

3D Protein Structure

Visualize protein structures in 3D. Users can display sequence conservation score on a structure and highlight experimentally determined epitopes as well.

Key Highlights:

- · Visualize protein structure in 3D
- Display sequence conservation heat map on the structure
- Highlight sequence features (epitopes, etc.)
- · Download highlighted protein structure image

Tutorial

Announcements

View 3D Structure

Start Search

Community Spotlight

View Archive

We would like to recognize the contribution of all our

Comparative Analysis of MERS-CoV Sequences

We have recently completed a comparative genomics analysis of Middle East Respiratory Syndrome Coronavirus (MERS-CoV) whole genome sequences, with implications for viral evolution, performed using the suite of bioinformatics tools available in ViPR. Also see a digest of recent events concerning MERS-CoV here.

What's New with Flu

An article in Cell says that a single amino acid change enhances H7N9 binding to lung receptors, but no new human H7N9 cases reported in a month. WHO implements a new 4phase pandemic alert system, and issues a new H7N9 risk assessment. Find these and other flu developments in the latest IRD Influenza Digest (View Archive).

And be sure to see a sequence analysis with implications for H7N9 evolution, carried out by IRD scientists using IRD comparative genomics analysis tools.

Live Demo









Avian and non-human mammalian surveillance data

www.fludb.org

SUBMIT DATA

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Virus phenotypic characteristics

Host Factor Data (Prototype)

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Highlights 3D Protein Structure Visualize protein structures in 3D. Users can display sequence conservation score on a structure and highlight experimentally determined epitopes as well. Key Highlights: · Visualize protein structure in 3D Display sequence conservation heat map on the structure Highlight sequence features (epitopes, etc.) · Download highlighted protein structure image View 3D Structure Start Search Tutorial

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Host Factor Experiments

Listed below are experiments from NIAID funded studies of viral infection. A given study may be divided into multiple experiments based commonly on the type of biological measurement.

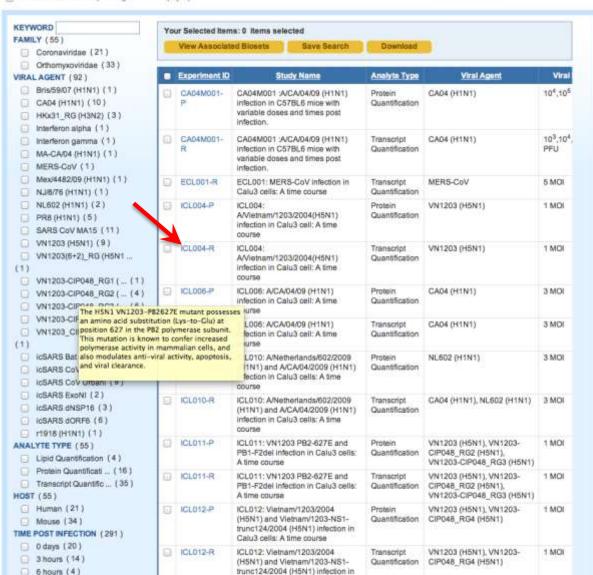
Search Help

- Keyword Search uses an exact match to fields Entrez Gene ID. Gene Accession, Symbol. Product Name.
- Selection of checkboxes will use a "or" for checkboxes in a category and "and" for groups of categories.

Displaying 55 of 55 records.

Data sorted by Experiment ID in ascending order

Select all 55 records (including those not displayed)



- 35 transcriptomic, 16 proteomic, 4 lipidomic experiments
- 2845 experiment samples
- 590 biosets
- 24 viral (flu, SARS, MERS) and 2 non-viral agents



Host Factor Experiment [ICLoo4-R]

Search Host Factor Experiments

Result Matrix

Previous Experiment Browse Next Experiment

Experiment Information | Experiment Sample Summary | Host Factor Bioset Information | Host Factor Bioset Summary | Host Factor Results

▲ Experiment Information

Study Name:	ICL004: A/Vietnam/1203/2004(H5N1) infection in Calu3 cell: A time course
Experiment Name:	ICL004-R: Calu-3 infections with A/Vietnam/1203-CIP048_RG1/2004(H5N1)
PI:	Michael Katze
Point of Contact:	Lynn Law, University of Washington, Department of Microbiology, Seattle, WA, gllaw@u.washington.edu; Michael Katze, University of Washington, Department of Microbiology, Seattle, WA, honey@u.washington.edu
Experiment Type:	Transcript Quantification
Measurement Technique:	Array, Agilent Techno_G4112F, geo GPL6480-26599_Agilent-014850 Whole Human Genome Microarray 4x44K G4112F (Probe Name version)
Description:	Purpose: To obtain samples from Calu-3 cells infected with A/Vietnam/1203/2004 (H5N1) for both transcriptional and proteomics analyses. Details: Time Points = 0*, 3*, 7, 12, 18*, and 24* hours post infection (*note: for proteomics the 0 and 3 hour samples were pooled and the 18 and 24h samples were pooled); Done in triplicate for both RNA and Protein; Triplicates are defined as 3 different wells, plated at the same time using the same cell stock for all replicates; Time matched mocks done in triplicate from same cell stock as rest of samples; Inoculation medium for mock infection was the same as the medium used for virus infection. Infection done at an MOI of 1.
Experimenters:	Yoshi Kawaoka, Dick Smith, Michael Katze, Shannon McWeeney, Katrina Waters
PubMed ID:	21865398 ଜ୍ର 22074594 ଜ୍ର
Conditional Variables:	Time points
Host Species Used:	Human
Geo Accession:	GSE28166 원
Protocols Used:	T003.0P _Nanochip_Bioanalyzer_protocol TCL001.0P - Preparation of Samples from Calu-3 cells for Isolation of RNA T002.0P Qiagen_RNeasy_Mini_Protocol SCL002.0P - Maintenance, Plating and Virus Infection of Calu-3 cells T006.0P Operating_the_Agilent_Microarray_Scanner T005.0P 4X44K_Hyb_only_protocol T004.1P cRNA_Probe_Synthesis
Viral Agent:	VN1203 (H5N1) High level of pathogenicity

▲ Experiment Sample Summary

# Animal Subjects	Subject Species	Subject Strain/Line	Strain/Line Characteristics	# Biological Samples	Biological Sample Source	Viral Agent	Viral Dose	Time Post Infection	# Experiment Samples
1	Homo sapiens	Calu-3 cells		36	Cell Line	VN1203 (H5N1)	1 MOI	0,3,7,12,18,24 hours	36



Pub Med.gov # 22074594[uid] PubMed US National Library of Medicine National Institutes of Health

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8 Search

Help

Display Settings: V Abstract

BMC Syst Biol. 2011 Nov 11;5:190, doi: 10.1186/1752-0509-5-190,

Conserved host response to highly pathogenic avian influenza virus infection in human cell culture, mouse and macaque model systems.

McDermott JE, Shankaran H, Eisfeld AJ, Belisle SE, Neuman G, Li C, McWeeney S, Sabourin C, Kawaoka Y, Katze MG, Waters KM.

Author information

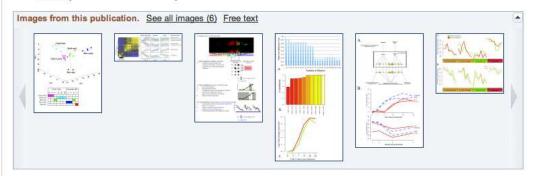
Abstract

BACKGROUND: Understanding host response to influenza virus infection will facilitate development of better diagnoses and therapeutic interventions. Several different experimental models have been used as a proxy for human infection, including cell cultures derived from human cells, mice, and non-human primates. Each of these systems has been studied extensively in isolation, but little effort has been directed toward systematically characterizing the conservation of host response on a global level beyond known immune signaling cascades.

RESULTS: In the present study, we employed a multivariate modeling approach to characterize and compare the transcriptional regulatory networks between these three model systems after infection with a highly pathogenic avian influenza virus of the H5N1 subtype. Using this approach we identified functions and pathways that display similar behavior and/or regulation including the well-studied impact on the interferon response and the inflammasome. Our results also suggest a primary response role for airway epithelial cells in initiating hypercytokinemia, which is thought to contribute to the pathogenesis of H5N1 viruses. We further demonstrate that we can use a transcriptional regulatory model from the human cell culture data to make highly accurate predictions about the behavior of important components of the innate immune system in tissues from whole organisms.

CONCLUSIONS: This is the first demonstration of a global regulatory network modeling conserved host response between in vitro and in vivo models.

PMID: 22074594 [PubMed - indexed for MEDLINE] PMCID: PMC3229612 Free PMC Article



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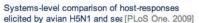


BMC Syst Biol Free full text

Save items



Related citations in PubMed



Highly pathogenic avian influenza H5N1 viruses elicit an attenuated type i interferor [J Virol. 2007]

Host regulatory network response to infection with highly pathogenic H5N1 avian [J Virol. 2011]

Review New strategies for the development of H5N1 subtype influenza vaccine [BioDrugs. 2011]

Review Innate immune responses to influenza A H5N1: friend or foe? [Trends Immunol. 2009]

See reviews...

See all...



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A semiautomated framework for integrating expert knowledge into diseas [Dis Markers, 2013]

Old world monkeys and new age science: the evolution of nonhuman primate sy [ILAR J. 2013]

A network integration approach to predict conserved regulators related to [PLoS One. 2013]

See all...

O comments

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Previous Experiment Browse Next Experiment

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Experiment Sample Details [ICL004-R]

Download

Previous Experiment Browse Next Experiment

Row	Experiment Sample User- Defined ID	Subject Species Name	Subject Strain/Line	Biological Sample Source	Treatment 1	Treatment 1 Dose	Treatment 1 Duration	Biological Sample Type	Source
1	251485048466_1_1_ES	Homo sapiens	Calu-3 cells	Cell Line	VN1203 (H5N1)	1 MOI	0 hours	RNA	GSM697564 &
2	251485048495_1_1_ES	Homo sapiens	Calu-3 cells	Cell Line	VN1203 (H5N1)	1 MOI	0 hours	RNA	GSM697566 &
3	251485048496_1_2_ES	Homo sapiens	Calu-3 cells	Cell Line	VN1203 (H5N1)	1 MOI	0 hours	RNA	GSM697583 &
4	251485048496_1_4_ES	Homo sapiens	Calu-3 cells	Cell Line	VN1203 (H5N1)	1 MOI	0 hours	RNA	GSM697582 ₫
5	251485048497_1_2_ES	Homo sapiens	Calu-3 cells	Cell Line	VN1203 (H5N1)	1 MOI	0 hours	RNA	GSM697584 &
6	251485048498_1_3_ES	Homo sapiens	Calu-3 cells	Cell Line	VN1203 (H5N1)	1 MOI	0 hours	RNA	GSM697565 &
7	251485048465_1_1_ES	Homo sapiens	Calu-3 cells	Cell Line	VN1203 (H5N1)	1 MOI	3 hours	RNA	GSM697568 ₫
8	251485048468_1_2_ES	Homo sapiens	Calu-3 cells	Cell Line	VN1203 (H5N1)	1 MOI	3 hours	RNA	GSM697569 &
9	251485048468_1_3_ES	Homo sapiens	Calu-3 cells	Cell Line	VN1203 (H5N1)	1 MOI	3 hours	RNA	GSM697567 &
10	251485048469_1_3_ES	Homo sapiens	Calu-3 cells	Cell Line	VN1203 (H5N1)	1 MOI	3 hours	RNA	GSM697585 ₫
11	251485048497_1_4_ES	Homo sapiens	Calu-3 cells	Cell Line	VN1203 (H5N1)	1 MOI	3 hours	RNA	GSM697586 &
12	251485048498_1_1_ES	Homo sapiens	Calu-3 cells	Cell Line	VN1203 (H5N1)	1 MOI	3 hours	RNA	GSM697587 &
13	251485048467_1_1_ES	Homo sapiens	Calu-3 cells	Cell Line	VN1203 (H5N1)	1 MOI	7 hours	RNA	GSM697570 &
14	251485048469_1_2_ES	Homo sapiens	Calu-3 cells	Cell Line	VN1203 (H5N1)	1 MOI	7 hours	RNA	GSM697589 &
15	251485048495_1_4_ES	Homo sapiens	Calu-3 cells	Cell Line	VN1203 (H5N1)	1 MOI	7 hours	RNA	GSM697571 ₫
16	251485048497_1_3_ES	Homo sapiens	Calu-3 cells	Cell Line	VN1203	1 MOI	7 hours	RNA	GSM697590 ₫



GEO Publications FAQ MISAME Email GEO

Query DataSets for GSM697564

NCBI - GEO - Accession Display (I)

Not logged in | Logn [7]

GEO heig: Mouse over screen elements for information.

Scope: Self : Format: HTML : Amount: Quick : GEO accession: GSM697564

Sample G5M697564

Status Public on Sep.02, 2011

Title Mock OH 1 Sample type RNA.

calu3, mock, 0H Source name Organism Homo sapiens Characteristics cell line: Calu-3

cell type: lung adenocarcinoma

infection: mock infection duration: 0h

Treatment protocol

For RNA isolation, Calu-3 cells were seeded in 6-well plates (1 x 10^6 cells/well) two days prior to infection. Immediately preceding infection,

monolayers were washed twice with DF12 supplemented with 0.3% bovine serum albumin (DF12-BSA), and inoculated with VN1203 (multiplicity of infection [MOI] of 1 plaque forming unit per cell) in DF12-BSA for 50 minutes at 37°C. Mock-infected controls were inoculated with DF12-BSA only. Following inoculation, monolayers were washed once with DF12-BSA and incubated in DF12-BSA containing 0.5 µg/ml of TPCK-treated trypsin

(Worthington Biochemical Corporation, Lakewood, NJ) for the times indicated. Growth protocol Calu-3 cells, a human lung adenocarcinoma cell line, were kindly provided by

Dr. Raymond Pickies (University of North Carolina, Chapel Hill, NC) and were maintained in a 1:1 mixture of Dulbecco's modified Eagle's medium and Ham's F12 nutrient medium (DF12; Invitrogen, Carlsbad, CA) supplemented with 10% fetal bovine serum. All cells were grown at 37°C in an atmosphere

of 5% CO2, with an antibiotic/antimycotic mixture (Invitrogen).

Extracted molecule

Extraction protocol At 0, 3, 7, 12, 18 and 24 hours post-infection (hpl), triplicate wells of mockinfected and VN1203-infected Calu-3 monolayers were washed with 5 ml cold

phosphate-buffered saline (PBS) and lysed directly with 1 ml of TRIzol (Invitrogen), according to the manufacturer's recommendation. The resulting lysates were stored at -80°C until further processing. All TRIzol lysates were processed simultaneously: they were phase-separated, and RNA was isolated from the aqueous phase (diluted 2 fold with RLT buffer) using Qiagen RNeasy Mini columns and the manufacturer's recommended protocol (Qiagen Inc., Valencia, CA). RNA quality was assessed on an Agilent 2100 Bioanalyzer using the nanochip format, and only intact RNA was used for quantitative real-time

PCR (qPCR) and microarray analyses.

Label Cy3

Scan protocol

Description

The Agilent One-Color Microarray-Based Gene Expression Analysis Protocol Label protocol

was followed for all processing steps, including Cy3-cDNA probe preparation.

Hybridization protocol The Agilent One-Color Microarray-Based Gene Expression Analysis Protocol

was followed for all processing steps, including hybridization and array

Dry slides were scanned on an Agilent DNA microarray scanner (Model

G2505B) using the XDR setting.

251485048466_1_1

Mock host response 0H.

Data processing Raw images were analyzed using the Agilent Feature Extraction software

(version 9.5.3.1) and the GE1-v5_95_Feb07 extraction protocol. Data were

normalized using RMA.

Submission date Mar 24, 2011 Last update date Sep 02, 2011 Contact name Armand Bankhead III

Organization name
Oregon Health and Science University

Departament of Medical Informatics and Clinical Epidemiology Department

Street address 3181 SW Sam Jackson Park Rd.

City Portland State/province OR ZIP/Postal code 97080 Country USA



▲ Host Factor Bioset Information

Bioset Type:	Differentially expressed genes
Protocol Used:	M001.0P - Statistical Protocol - Normalization, QAQC and Differential Expression Analysis for Agilent Arrays
Description:	This is a results matrix that lists all DE genes compared to time matched mock samples for Calu-3 cell samples infected with A/Vietnam/1203-CIP048_RG1/2004(H5N1) at an MOI of 1. Time points are 0, 3, 7, 12, 18, and 24 h post infection. Differential Expression Criteria: passes Agilent QC flag and FC > 1.5 and q-value < .05.
Analysis Method:	Agilent Preprocess 44x4K/LIMMA
Normalization Method:	quantile
Differential Expression Criteria Used:	passes Agilent QC flag and log2 FC > 1.5 and q-value < .05

▲ Host Factor Bioset Summary

Bioset Name	Host Factors	Viral Agent	Viral Dose	Time Post Infection	Strain/Line	Host
ICL004-R_0	0	VN1203 (H5N1)	1 MOI	0 hours	Calu-3 cells	Human
ICL004-R_3	0	VN1203 (H5N1)	1 MOI	3 hours	Calu-3 cells	Human
ICL004-R_7	5,277	VN1203 (H5N1)	1 MOI	7 hours	Calu-3 cells	Human
ICL004-R_12	13,030	VN1203 (H5N1)	1 MOI	12 hours	Calu-3 cells	Human
ICL004-R_18	12,944	VN1203 (H5N1)	1 MOI	18 hours	Calu-3 cells	Human
ICL004-R_24	17,324	VN1203 (H5N1)	1 MOI	24 hours	Calu-3 cells	Human







▲ Host Factor Bioset Patterns

Your Selected Items: 0 items selected

Host Factor Results

Download

The Host Factor Results button will create a union of the selected items and build a table below Host Factor Bioset Patterns.

Significant Host	Factors as	Compared to	Control
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"+" up-regulated, "-" down-regulated, blank: no change

SYMBOL Find expression pattern:

Use comma to separate multiple entries. Ex. DDX58

Find

Your search returned 55 records.

Data sorted by Number of Probes in descending order

•	Pattern	Virus Time Post Infection Viral Dose Strain/Line	VN1203 (H5N1), 0 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 3 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 7 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 12 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 18 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 24 hours, 1 MOI, Calu-3 cells
		Host Factors	0	0	□5277	<u> </u>	<u>12944</u>	□ 17324
	0,0,0,+,+,+	2921				+	+	+
	0,0,0,-,-,-	2764				-	-	-
	+,0,0,0,0,0	2105						+
	0,0,0,0,0,-	1974						-
	0,0,-,-,-,-	1643			-	-	-	-
	0,0,0,0,-,-	1621					-	-
	0,0,+,+,+,+	1094			+	+	+	+
	0,0,0,0,+,+	931					+	+
	0,0,+,0,0	919				+		
	0,0,-,0,0	746				-		
	0,0,0,-,0,-	487				-		-
	0,0,0,+,0,+	452				+		+
	0,0,+,0,0	433			+			
	0,0,0,+,+,0	433				+	+	
	0,0,0,-,-,0	402				-	-	
	0,0,-,0,0	275			-			
	0,-,0,0,0,-	230					-	

Host Factor Experiments

Listed below are experiments from NIAID funded studies of viral infection. A given study may be divided into multiple experiments based commonly on the type of biological measurement.

Search Help:

- · Keyword Search uses an exact match to fields Entrez Gene ID, Gene Accession, Symbol, Product Name.
- · Selection of checkboxes will use a "or" for checkboxes in a category and "and" for groups of categories.

Displaying 55 of 55 records.

Data sorted by Experiment ID in ascending order

KEYV	VORD	You	ur Selected Item	s: 2 items selected Deselect All			
FAMI	LY (55)		View Associate	ed Blosets Save Search	Download		
	Coronaviridae (21)		TICH ASSOCIATI	Sure Startin	Dominoud		
	Orthomyxoviridae (33)	Same I	Para la	O	The state of the state of		30-1
VIRA	LAGENT (92)		Experiment ID	Study Name	Analyte Type	Viral Agent	Viral
0	Bris/59/07 (H1N1) (1)		CA04M001-	CA04M001 :A/CA/04/09 (H1N1)	Protein	CA04 (H1N1)	10 ⁴ ,10 ⁵
	CA04 (H1N1) (10)		P	infection in C57BL6 mice with variable doses and times post infection.	Quantification		
U	HKx31_RG (H3N2) (3)	1					
	Interferon alpha (1)		CA04M001-	CA04M001 :A/CA/04/09 (H1N1)	Transcript	CA04 (H1N1)	10 ³ ,10 ⁴ ,
	Interferon gamma (1)	1 -	R	infection in C57BL6 mice with Quantification		CA04 (HTN1)	PFU
	MA-CA/04 (H1N1) (1)	1		variable doses and times post			
	MERS-CoV (1) Mex/4482/09 (H1N1) (1) NJ/8/76 (H1N1) (1)	_		infection.			
		0	ECL001-R	ECL001: MERS-CoV infection in Calu3 cells: A time course	Transcript Quantification	MERS-CoV	5 MOI
	NL602 (H1N1) (2)		ICL004-P	ICL004:	Protein	VN1203 (H5N1)	1 MOI
0	PR8 (H1N1) (5)			A/Vietnam/1203/2004(H5N1) Quantification infection in Calu3 cell: A time course	Quantification		
	SARS CoV MA15 (11)	1					
	VN1203 (H5N1) (9)	()	ICL004-R	ICL004:	Transcript Quantification	VN1203 (H5N1)	1 MOI
	VN1203(6+2)_RG (H5N1	1.00		A/Vietnam/1203/2004(H5N1)			
(1)		1		infection in Calu3 cell: A time			
	VN1203-CIP048_RG1 ((1)	JAN.	Market State	course			800000000
	VN1203-CIP048_RG2 ((4)		ICL006-P	ICL006: A/CA/04/09 (H1N1) infection in Calu3 cell: A time	Protein Quantification	CA04 (H1N1)	3 MOI
	VN1203-CIP048_RG3 ((6)			course	Quantification		
	VN1203-CIP048_RG4 ((4)	d	ICL006-R	ICL006: A/CA/04/09 (H1N1)	Transcript	CA04 (H1N1)	3 MOI
	VN1203_CIP048_RG1(H5			infection in Calu3 cell: A time	Quantification	PROCESS OF MARKETS (A)	3 11101
(1)				course			
	icSARS Bat SRBD (6)		ICL010-P	ICL010: A/Netherlands/602/2009	Protein	NL602 (H1N1)	3 MOI
	icSARS CoV (1) icSARS CoV Urbani (9)			(H1N1) and A/CA/04/2009 (H1N1) infection in Calu3 cells: A time	Quantification		
	icSARS ExoNI (2)		101.040.0	course		0.104 (1.141) 111 000 (1.111)	0.1101
	icSARS dNSP16 (3)	☑	ICL010-R	ICL010: A/Netherlands/602/2009 (H1N1) and A/CA/04/2009 (H1N1)	Transcript Quantification	CA04 (H1N1), NL602 (H1N1)	3 MOI
	icSARS dORF6 (6)			infection in Calu3 cells: A time	Saurimoudon		
	r1918 (H1N1) (1)			course			

Host Factor Biosets

Search Help:

- . Keyword Search uses an exact match to fields Entrez Gene ID, Gene Accession, Symbol, Product Name.
- . Selection of checkboxes will use a "or" for checkboxes in a category and "and" for groups of categories.
- · If no checkbox is shown for a bioset, no host factors are currently available.

Displaying 22 of 22 records.

Data sorted by Bioset Name in ascending order

Select all 22 records (including those not displayed)

	WORD	Yo	ur Selected Items: 2 Items se	lected	Deselect All				
	ERIMENT NAME (22)		Patterns Save Sear	ch	Set Operations	V	Host Factor Resu	.5	
	CA04M001-P: A/CA/04/ (0)	-	Download		Find shared facto	rs (Interse	et AND)		
	CA04M001-R: Mouse in (0)		DOMINOU		Find all factors (U	- A			
(0)	ECL001-R: MERS-CoVI		Bloset Name	Host	- 47	77 <u>12</u>		tion Strain/Line	Host
(0)	ICL004-P: A/Vietnam/ (0)		ICL006 Ohr H1N1	0	Find unique factor	s (XOR)	<u> </u>	Calu-3	Human
	ICL004-R: Calu-3 inf (0)	1	ICC000_UIII_HTINT	U	Find exclusive fac	ctors (Subt	ract)	cells	numan
	ICL006-P: A/CA/04/09 (0)		ICL006 12hr H1N1	1141	CA04	3 MOI	12 hours	Calu-3	Human
1	ICL006-R; Calu-3 inf (9)				(H1N1)			cells	
	ICL010-P: A/Netherla (0)		ICL006_18hr_H1N1	1948	CA04	3 MOI	18 hours	Calu-3	Human
V	ICL010-R: A/Netherla (13)				(H1N1)			cells	
	ICL011-P: VN1203 PB2 (0)	◙	ICL006_24hr_H1N1	1546	CA04 (H1N1)	3 MOI	24 hours	Calu-3 cells	Human
	ICL011-R: VN1203 PB2 (0)	0	ICL006_30hr_H1N1	1326	CA04	3 MOI	30 hours	Calu-3	Human
	ICL012-P: Vietnam/12 (0)				(H1N1)			cells	
	ICL012-R; Vietnam/12 (0)	0	ICL006_36hr_H1N1	1255	CA04	3 MOI	36 hours	Calu-3	Human
	IM001-P: A/Vietnam/1 (0)				(H1N1)			cells	
	IM001-R: Mouse infec (0)		ICL006_3hr_H1N1	0	CA04 (H1N1)	3 MOI	3 hours	Calu-3 cells	Human
	IM002-R: Influenza A (0)		101 000 405 - 11414	4400		0.1101	40.1		110.02150
	IM004-P: VN1203 HA a (0)		ICL006_48hr_H1N1	1190	CA04 (H1N1)	3 MOI	48 hours	Calu-3 cells	Human
	IM004-R: Mouse infec (0)		ICL006 7hr H1N1	360	CA04	3 MOI	7 hours	Calu-3	Human
	IM005-P: Vietnam/120 (0) IM005-R: Vietnam/120 (0)		102000_1111_111111	000	(H1N1)	o mor	1 110010	cells	. Toman
	IM006A-P: Vietnam/120 (0)		ICL010_Cal_0h_array_DE	0	CA04	3 MOI	0 hours	Calu-3	Human
	IM006A-R: Vietnam/12 (0)		a con on on ill am		(H1N1)			cells	
0	IM006B-P: Vietnam/12 (0)	0	ICL010_Cal_12h_array_DE	459	CA04 (H1N1)	3 MOI	12 hours	Calu-3 cells	Human
	IM006B-R: Vietnam/12 (0)	Ø	ICL010_Cal_24h_array_DE	756	CA04	3 MOI	24 hours	Calu-3	Human
	IM007-P: Vietnam/120 (0)	1207.5			(H1N1)			cells	
	IM007-R: Vietnam/120 (0)		ICL010_Cal_48h_array_DE	734	CA04	3 MOI	48 hours	Calu-3	Human
	IM009-R: Influenza A (0)			10040	(H1N1)			cells	24/10/2004
	IM010-R: Vietnam/120 (0)		ICL010_NL_0h_array_DE	0	NL602 (H1N1)	3 MOI	0 hours	Calu-3 cells	Human
	IM015-R: Influenza A (0)		ICL010_NL_12h_array_DE	105	NL602	3 MOI	12 hours	Calu-3	Human
	SBRI_AA: Transcripto (0)		IOLOTO_NL_1ZII_allay_DE	100	(H1N1)	3 IVIOI	12 Hours	cells	numan
	SBRI_LAE: BALF lipid (0)	0	ICL010 NL 18h array DE	226	NL602	3 MOI	18 hours	Calu-3	Human
	SBRI_LI: BALF lipido (0) SBRI_LN: BALF lipido (0)			-	(H1N1)			cells	. SATERIAL INC.
0	SBRI_LV: BALF lipido (0)	0	ICL010_NL_24h_array_DE	580	NL602	3 MOI	24 hours	Calu-3	Human
	CON_LY. DALI IIpido (0)				(H1N1)			cells	- Panaga-dan

Boolean Analysis Result for Derived Bioset

Search Help:

- Keyword Search uses an exact match to fields Entrez Gene ID, Gene Accession, Symbol, Product Name.
- · Selection of checkboxes will use a "or" for checkboxes in a category and "and" for groups of categories.

Displaying 200 of 502 records.

Data sorted by Gene Symbol, Probe Id in ascending order

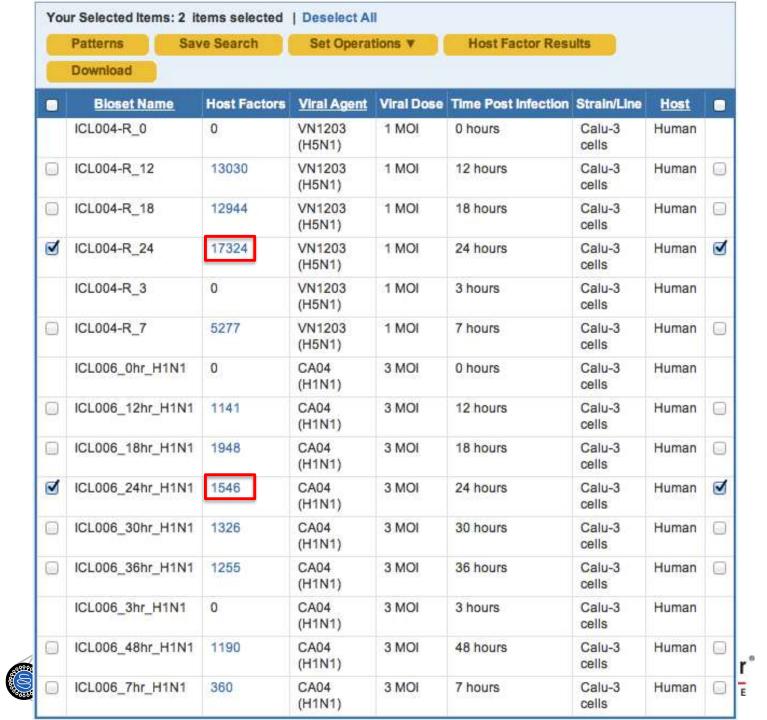
	7/05	Yo	ur Selected Items	: 0 items selecte	d				
Inter	TYPE		Set Operations V	Save Se	earch	Add to Working Set	Download		
	ERIMENT NAME (2)								
	CA04M001-P: A/CA/04/ (0)							ı,	ICL
	CA04M001-R: Mouse in (0)		Host Factor	Entrez Gene ID	Symbol	Genbank Accessio	n Name	lmm	oort L
(0)	ECL001-R: MERS-CoV i ICL004-P: AV/ietnam/ (0)		A_24_P263878	27	abl2	NM_007314	v-abl abelson murine leukemia viral oncogene homolog 2 (arg, abelson-	&	1.8
1 60	ICL004-R; Calu-3 inf (0)	_		122		2000 2000	related gene)	V-902	- 2
	ICL006-P: A/CA/04/09 (0)	(6)	A_24_P46130	55	acpp	NM_001099	acid phosphatase, prostate	Ġ.	-3
₫	ICL006-R: Calu-3 inf (1)	9	A_23_P1102	58	acta1	NM_001100	actin, alpha 1, skeletal muscle	<u>G</u>	2.6
✓	ICL010-P: A/Netherla (0) ICL010-R: A/Netherla (1)		A_23_P126363	11085	adam30	NM_021794	adam metallopeptidase domain 30	Ø	1.9
	ICL011-P: VN1203 PB2 (0)	0	A_23_P137786	54507	adamtsl4	AK023606	adamts-like 4	G	2.0
	ICL011-R: VN1203 PB2 (0) ICL012-P: Vietnam/12 (0)	0	A_23_P49816	55803	adap2	NM_018404	arfgap with dual ph domains 2	ß	1.
	ICL012-R: Vietnam/12 (0) IM001-P: A/Vietnam/1 (0)	0	A_23_P51787	270	ampd1	NM_000036	adenosine monophosphate deaminase 1 (isoform m)	Ø	4.2
	IM001-R: Mouse infec (0)	0	A_23_P159325	51129	angptl4	NM_139314	angiopoietin-like 4	₫.	3.
	IM002-R: Influenza A (0)		A_24_P357572	338699	ankrd42	NM_182603	ankyrin repeat domain 42	₫.	1.
	IM004-P: VN1203 HA a (0)		A 23 P155049	80830	apol6	NM 030641	apolipoprotein I, 6	P	2.
	IM004-R: Mouse infec (0)		A 24 P941167	80830	apol6	NM 030641	apolipoprotein I, 6	₫P	1.
	IM005-P: Vietnam/120 (0) IM005-R: Vietnam/120 (0)	0	A_24_P380061	83478	arhgap24	NM_001025616	rho gtpase activating protein 24	ď	-2
	IM006A-P: Vietnam/12 (0) IM006A-R: Vietnam/12 (0)	0	A_24_P414553	57561	arrdc3	NM_020801	arrestin domain containing	ď	1.9
	IM006B-P: Vietnam/12 (0) IM006B-R: Vietnam/12 (0)	0	A_23_P155265	142686	asb14	NM_130387	ankyrin repeat and socs box-containing 14	ď	3.
	IM007-P: Vietnam/120 (0) IM007-R: Vietnam/120 (0)	0	A_23_P302750	83858	atad3b	AB033099	atpase family, aaa domain containing 3b	ď	2.3
	IM009-R: Influenza A (0) IM010-R: Vietnam/120 (0)	0	A_23_P118894	79170	atad4	NM_024320	atpase family, aaa domain containing 4	Ø	-1
0	IM015-R: Influenza A (0) SBRI AA: Transcripto (0)	0	A_24_P405205	493	atp2b4	NM_001001396	atpase, ca++ transporting, plasma membrane 4	Ø	-1.
	SCL005-P: icSARS ORF	(3)	A 23 P53257	10677	avil	BX647344	advillin	₫.	2.5

	Genhank Access			ICL010_Cal_24h_array_DE ICL006_24hr_H1N1					
ř	Genbank Accessi	on Name	lmmp	ort Log2 FC	Q-Value	Log2 FC	P-Value		
	NM_007314	v-abl abelson murine leukemia viral oncogene homolog 2 (arg, abelson- related gene)	_E	1.8	1.3E-2	1.9	1.0E-3	0	
	NM_001099	acid phosphatase, prostate	₫P	-3.0	4.1E-3	-1.9	8.3E-4		
	NM_001100	actin, alpha 1, skeletal muscle	Ø.	2.6	1.6E-3	3.0	2.5E-3		
	NM_021794	adam metallopeptidase domain 30	&	1.9	2.4E-2	2.3	7.2E-3	0	
	AK023606	adamts-like 4	₫P	2.0	3.2E-2	1.9	4.4E-2		
	NM_018404	arfgap with dual ph domains 2	G.	1.5	6.9E-3	1.7	9.3E-3		
	NM_000036	adenosine monophosphate deaminase 1 (isoform m)	g.	4.2	2.1E-3	4.0	6.4E-3		
	NM_139314	angiopoietin-like 4	₫P	3.5	1.5E-3	1.9	5.3E-4		
	NM_182603	ankyrin repeat domain 42	蛩	1.5	4.2E-2	1.8	9.8E-3		
	NM_030641	apolipoprotein I, 6	g.	2.8	4.9E-3	1.5	8.7E-3		
	NM_030641	apolipoprotein I, 6	₫.	1.7	3.5E-3	2.2	5.0E-3		
	NM_00 <mark>1</mark> 025616	rho gtpase activating protein 24	G	-2.1	3.6E-3	-1.5	2.0E-2		
	NM_020801	arrestin domain containing 3	g.	1.9	2.2E-3	1.5	3.6E-3	0	
	NM_130387	ankyrin repeat and socs box-containing 14	₽.	3.1	3.8E-3	2.9	2.1E-2		
	AB033099	atpase family, aaa domain containing 3b	ক্র	2.3	5.0E-3	1.7	4.1E-2	0	
	NM_024320	atpase family, aaa domain containing 4	g.	-1.9	2.7E-3	-2.6	2.7E-4		
	NM_001001396	atpase, ca++ transporting, plasma membrane 4	<u>G</u>	-1.5	2.1E-3	-2.0	5.3E-3		
	BX647344	advillin	ন্ত্ৰ	2.5	5.3E-4	1.7	1.6E-3		
	NM_004655	axin 2	Ø.	-1.9	3.2E-3	-1.5	1.6E-2		
)	AK022379	beta-2-microglobulin	ক্র	2.9	2.0E-3	2.9	1.9E-3		
	NM_138456	basic leucine zipper transcription factor, atf-like 2	굡	4.2	7.0E-4	3.3	3.1E-4	0	





0	Experiment ID	Study Name	Analyte Type	Viral Agent	Viral
0	CA04M001- P	CA04M001 :A/CA/04/09 (H1N1) infection in C57BL6 mice with variable doses and times post infection.	Protein Quantification	CA04 (H1N1)	10 ⁴ ,10 ⁵
	CA04M001- R	CA04M001 :A/CA/04/09 (H1N1) infection in C57BL6 mice with variable doses and times post infection.	Transcript Quantification	CA04 (H1N1)	10 ³ ,10 ⁴ , PFU
	ECL001-R	ECL001: MERS-CoV infection in Calu3 cells: A time course	Transcript Quantification	MERS-CoV	5 MOI
0	ICL004-P	ICL004: A/Vietnam/1203/2004(H5N1) infection in Calu3 cell: A time course	Protein Quantification	VN1203 (H5N1)	1 MOI
Ø	ICL004-R	ICL004: A/Vietnam/1203/2004(H5N1) infection in Calu3 cell: A time course	Transcript Quantification	VN1203 (H5N1)	1 MOI
0	ICL006-P	ICL006: A/CA/04/09 (H1N1) infection in Calu3 cell: A time course	Protein Quantification	CA04 (H1N1)	3 MOI
⋖	ICL006-R	ICL006: A/CA/04/09 (H1N1) infection in Calu3 cell: A time course	Transcript Quantification	CA04 (H1N1)	3 MOI
0	ICL010-P	ICL010: A/Netherlands/602/2009 (H1N1) and A/CA/04/2009 (H1N1) infection in Calu3 cells: A time course	Protein Quantification	NL602 (H1N1)	3 MOI
	ICL010-R	ICL010: A/Netherlands/602/2009 (H1N1) and A/CA/04/2009 (H1N1) infection in Calu3 cells: A time course	Transcript Quantification	CA04 (H1N1), NL602 (H1N1)	3 MOI



		Yo	ur Selected Items	: 0 items selecte	d				
	TYPE		Set Operations V	Save Se	arch A	Add to Working Set	Download		
Inter	SECT ERIMENT NAME (2)								
	CA04M001-P: A/CA/04/ (0)	Ĩ.							ICLO
0	CA04M001-R: Mouse in (0)		Host Factor	Entrez Gene ID	Symbol	Genbank Accessio	n Name	lmmt	ottog2
0	ECL001-R: MERS-CoV I		A_23_P9415	48	aco1	NM_002197	aconitase 1, soluble	g.	-2.6
(0)			A_23_P9416	48	aco1	NM_002197	aconitase 1, soluble	g.	-14.7
	ICL004-P: A/Vietnam/ (0)		A 23 P317756	6296	MATERIAL CONTRACTOR OF THE PROPERTY OF THE PRO	NM 202000		- North	-2.3
V	ICL004-R: Calu-3 inf (1)		A_23_P317750	0290	acsm3	NM_202000	acyl-coa synthetase medium-chain	₫.	-2.3
0	ICL006-P: A/CA/04/09 (0)	1					family member 3		
1	ICL006-R: Calu-3 inf (1)	0	A_23_P1102	58	acta1	NM_001100	actin, alpha 1,	g.	2.6
0	ICL010-P: A/Netherla (0)					5.75	skeletal muscle		
	ICL010-R: A/Netherla (0)		A_23_P137786	54507	adamtsl4	AK023606	adamts-like 4	E	12.6
0	ICL011-P: VN1203 PB2 (0)		A_23_P49816	55803	adap2	NM_018404	arfgap with dual ph	&	1.5
	ICL011-R: VN1203 PB2 (0)					MATTAWA AND SALES	domains 2		
0	ICL012-P: Vietnam/12 (0)	0	A_23_P126313	55811	adcy10	NM_018417	adenylate cyclase	ď.	19.6
	ICL012-R: Vietnam/12 (0)	-		1999	2010	1111 000001	10 (soluble)		
	IM001-P: A/Vietnam/1 (0)		A_23_P145024	154	adrb2	NM_000024	adrenergic, beta-2-, receptor, surface	중	-1.9
	IM001-R: Mouse infec (0)	0	A_23_P76823	122622	adssl1	NM_199165	adenylosuccinate	r de	-1.7
0	IM002-R: Influenza A (0)	0	A_25_F70025	122022	aussii	MM_133103	synthase like 1	M	-1.7
	IM004-P: VN1203 HA a (0)	0	A_23_P214897	9590	akap12	NM_144497	a kinase (prka)	_©	2.7
	IM004-R: Mouse infec (0)	-					anchor protein 12		
	IM005-P: Vietnam/120 (0)		A_23_P257971	1645	akr1c1	NM_001353	aldo-keto reductase	₽.	-1.7
	IM005-R: Vietnam/120 (0)						family 1, member c1		
	IM006A-P: Vietnam/12 (0)	1					(dihydrodiol dehydrogenase 1;		
	IM006A-R: Vietnam/12 (0)	1					20-alpha (3-alpha)-		
	IM006B-P: Vietnam/12 (0)	1					hydroxysteroid		
0	IM006B-R: Vietnam/12 (0)	-		O LANGUAGO LA			dehydrogenase)	N/A	
	IM007-P: Vietnam/120 (0)		A_24_P220947	1645	akr1c1	NM_001353	aldo-keto reductase family 1, member c1	G.	-8.0
0	IM007-R: Vietnam/120 (0)						(dihydrodiol		
	IM009-R: Influenza A (0)						dehydrogenase 1;		
0	IM010-R: Vietnam/120 (0)						20-alpha (3-alpha)- hydroxysteroid		
	IM015-R: Influenza A (0)						dehydrogenase)		
-	CONTRACTOR OF THE PARTY OF THE						The second of th		

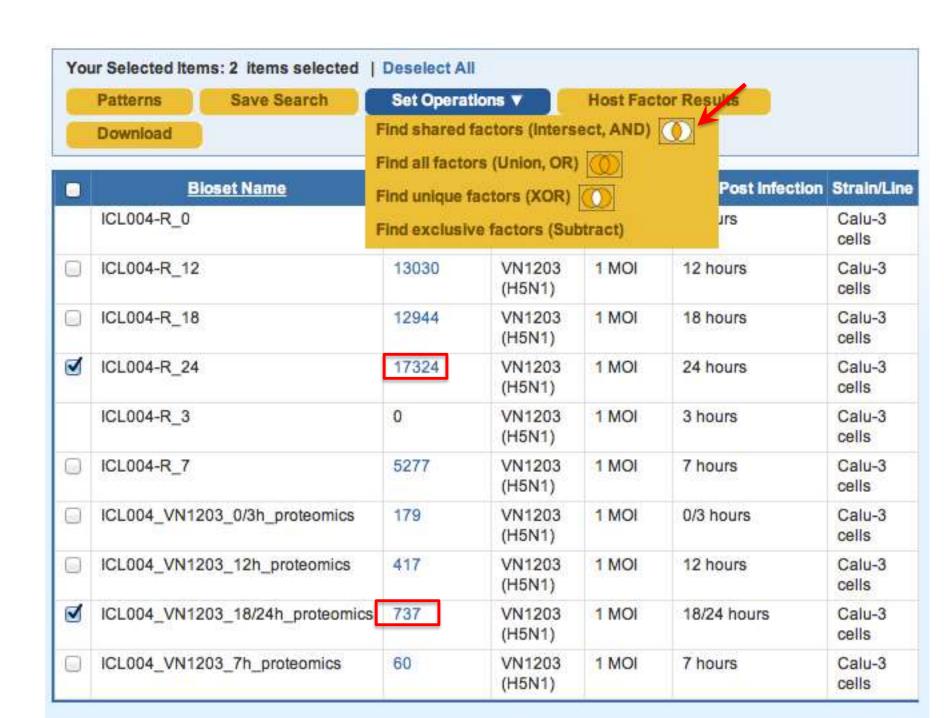
Your Selected Items: 2	2 items selected	Deselect All
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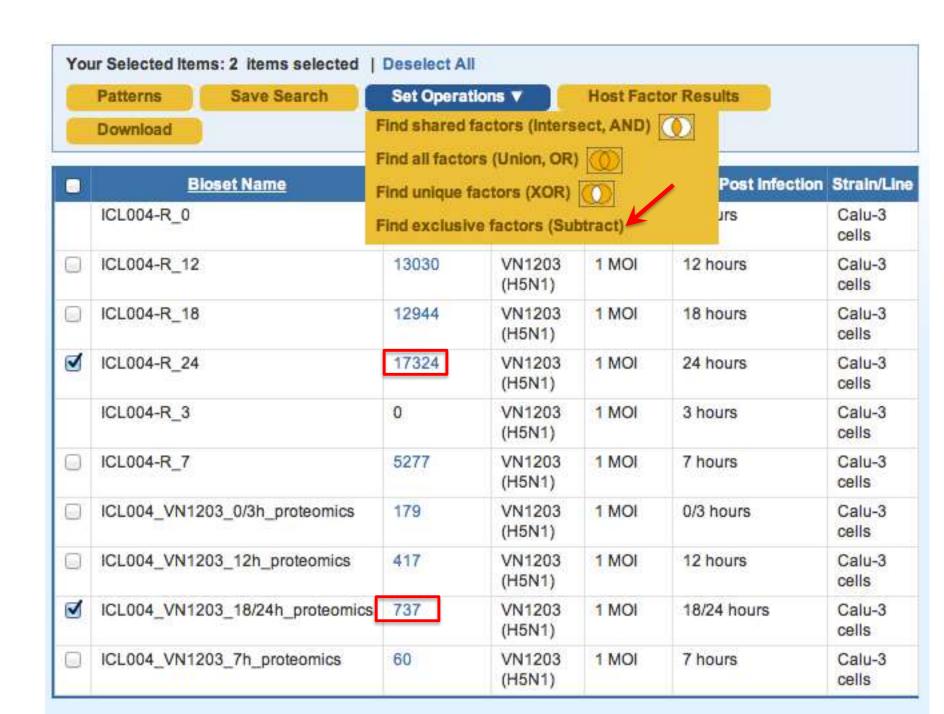
View Associated Biosets

Save Search

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	Experiment ID	Study Name	Analyte Type	Viral Agent	Viral
	CA04M001- P	CA04M001 :A/CA/04/09 (H1N1) infection in C57BL6 mice with variable doses and times post infection.	Protein Quantification	CA04 (H1N1)	10 ⁴ ,10 ⁵
0	CA04M001- R	CA04M001 :A/CA/04/09 (H1N1) infection in C57BL6 mice with variable doses and times post infection.	Transcript Quantification	CA04 (H1N1)	10 ³ ,10 ⁴ PFU
	ECL001-R	ECL001: MERS-CoV infection in Calu3 cells: A time course	Transcript Quantification	MERS-CoV	5 MOI
Ø	ICL004-P	ICL004: A/Vietnam/1203/2004(H5N1) infection in Calu3 cell: A time course	Protein Quantification	VN1203 (H5N1)	1 MOI
☑	ICL004-R	ICL004: A/Vietnam/1203/2004(H5N1) infection in Calu3 cell: A time course	Transcript Quantification	VN1203 (H5N1)	1 MOI
0	ICL006-P	ICL006: A/CA/04/09 (H1N1) infection in Calu3 cell: A time course	Protein Quantification	CA04 (H1N1)	3 MOI
	ICL006-R	ICL006: A/CA/04/09 (H1N1) infection in Calu3 cell: A time course	Transcript Quantification	CA04 (H1N1)	3 MOI
0	ICL010-P	ICL010: A/Netherlands/602/2009 (H1N1) and A/CA/04/2009 (H1N1) infection in Calu3 cells: A time course	Protein Quantification	NL602 (H1N1)	3 MOI
	ICL010-R	ICL010: A/Netherlands/602/2009 (H1N1) and A/CA/04/2009 (H1N1) infection in Calu3 cells: A time course	Transcript Quantification	CA04 (H1N1), NL602 (H1N1)	3 MOI





Boolean "Subtract"

Select one bioset from column A. Select one or more biosets from column B (hold down the shift key while selecting multiple values). Biosets selected from column B will be subtracted from the bioset selected in column A. The maximum sets that can be subtracted from column A is 4.

A

ICL004-R_24

ICL004_VN1203_18/24h_proteomics

В

ICL004-R_12 ICL004-R_18 ICL004-R_24 ICL004-R_7

Cancel

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		You	ur Selecte	ed Items: 0 Items selected			
	YPE		Set Opera	ations ▼ Save Search Add	d to Working Set	Download	
/linu XPE	RIMENT NAME (2)		10				
	CA04M001-P: A/CA/04/ (0)		HID	Bloset Name	Host Factor	Entrez Gene ID	Symbol
	CA04M001-R: Mouse in (0) ECL001-R: MERS-CoV i	0	347	ICL004_VN1203_18/24h_proteomics	GALE_HUMAN	2582	gale
0)	EGEOU I-R. MERG-GOV I		658	ICL004_VN1203_18/24h_proteomics	GSTM3_HUMAN	2947	gstm3
▼	ICL004-P: A/Vietnam/ (1) ICL004-R: Calu-3 inf (1)		726	ICL004_VN1203_18/24h_proteomics	PPIB_HUMAN	5479	ppib
	ICL006-P: A/CA/04/09 (0)		779	ICL004_VN1203_18/24h_proteomics	RS3_HUMAN	6188	rps3
	ICL006-R: Calu-3 inf (0) ICL010-P: A/Netherla (0)	0	803	ICL004_VN1203_18/24h_proteomics	RS27_HUMAN	6232	rps27
Ö	ICL010-R: A/Netherla (0)	0	980	ICL004_VN1203_18/24h_proteomics	CIRBP_HUMAN	1153	cirbp
	ICL011-P: VN1203 PB2 (0) ICL011-R: VN1203 PB2 (0)	0	1038	ICL004_VN1203_18/24h_proteomics	DOPD_HUMAN	1652	ddt
	ICL012-P: Vietnam/12 (0)		1039	ICL004_VN1203_18/24h_proteomics	DHX9_HUMAN	1660	dhx9
	ICL012-R: Vietnam/12 (0) IM001-P: A/Vietnam/1 (0)		1174	ICL004_VN1203_18/24h_proteomics	HNRPL_HUMAN	3191	hnrnpl
	IM001-R: Mouse infec (0)	0	1180	ICL004_VN1203_18/24h_proteomics	HSPB1_HUMAN	3315	hspb1
	IM002-R: Influenza A (0) IM004-P: VN1203 HA a (0)		1194	ICL004_VN1203_18/24h_proteomics	CYR61_HUMAN	3491	cyr61
	IM004-R: Mouse infec (0)	0	1250	ICL004_VN1203_18/24h_proteomics	COPD_HUMAN	372	arcn1
	IM005-P: Vietnam/120 (0) IM005-R: Vietnam/120 (0)	0	1275	ICL004_VN1203_18/24h_proteomics	AT5F1_HUMAN	515	atp5f1
	IM006A-P: Vietnam/12 (0) IM006A-R: Vietnam/12 (0)	0	1327	ICL004_VN1203_18/24h_proteomics	CPNS1_HUMAN	826	capns1
5	IM006B-P: Vietnam/12 (0)	0	1568	ICL004_VN1203_18/24h_proteomics	GNS_HUMAN	2799	gns
	IM006B-R: Vietnam/12 (0) IM007-P: Vietnam/120 (0)		1660	ICL004_VN1203_18/24h_proteomics	REXO4_HUMAN	57109	rexo4
	IM007-R: Vietnam/120 (0)	0	1707	ICL004_VN1203_18/24h_proteomics	IMB1_HUMAN	3837	kpnb1
	IM009-R: Influenza A (0) IM010-R: Vietnam/120 (0)	0	1855	ICL004_VN1203_18/24h_proteomics	MYPT1_HUMAN	4659	ppp1r12a
)	IM015-R: Influenza A (0)		1933	ICL004_VN1203_18/24h_proteomics	PA1B3_HUMAN	5050	pafah1b3
	SBRI_AA: Transcripto (0) SCL005-P: icSARS ORF		1945	ICL004_VN1203_18/24h_proteomics	PCNA_HUMAN	5111	pcna

Significant Host Factors as Compared to Control

"+" up-regulated, "-" down-regulated, blank: no change

Find expression pattern:

SYMBOL	

Find

Use comma to separate multiple entries. Ex. DDX58

Your search returned 55 records.

Data sorted by Number of Probes in descending order

	order										
-	Pattern	Virus Time Post Infection Viral Dose Strain/Line	VN1203 (H5N1), 0 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 3 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 7 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 12 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 18 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 24 hours, 1 MOI, Calu-3 cells			
		Host Factors	0	0	□ 5277	13030	12944	□ 17324			
	0,0,0,+,+,+	2921				+	+	+			
	0,0,0,-,-,-	2764				-	-	-			
	+,0,0,0,0,0	2105						+			
	0,0,0,0,0,-	1974						-			
	0,0,-,-,-	1643			-	-	-	-			
	0,0,0,0,-,-	1621					-	-			
	0,0,+,+,+,+	1094			+	+	+	+			
	0,0,0,0,+,+	931					+	+			
	0,0,0,+,0,0	919				+					
	0,0,-,0,0	746				-					
	0,0,0,-,0,-	487				-		-			
	+,0,+,0,0	452				+		+			
	0,0,+,0,0,0	433			+						
	0,0,0,+,+,0	433				+	+				
	0,0,0,-,-,0	402				-	-				
	0,0,-,0,0	275			-						
	0,0,0,0,-,0	230					-				
	0,0,-,-,0,0	222			-	-					
	0,0,-,-,0	195			-	-	-				
	0,0,+,0,0,+	192			+			+			
	0,0,-,-,0,-	180			-	-		-			
	0,0,0,0,+,0	152					+				
	0,0,+,0,+,+	133			+		+	+			
	0,0,0,+,0,-	115				+		-			
	0,0,+,+,0,0	110			+	+					
	0,0,-,0,-,-	102			-		-	-			
	0,0,-,0,0,-	102			-			-			

Displaying 500 of 1,094 records. Data sorted by Symbol, Host Factor ID in ascending order Select all 1,094 records (including those not displayed) Your Selected Items: 0 Items selected Add to Working Set Save Search Download Download Reactome Data Pathway View HOST FACTOR ID OR SYMBOL Find data for Find Reset Use comma to separate multiple entries. Ex. DDX58 Associated Bioset Information 1 ICL004: A/Vietnam/1203/2004(H5N1) infection in Calu3 cell: A time course Study Name **Experiment Name** ICL004-R: Calu-3 infections with A/Vietnam/1203-CIP048_RG1/2004(H5N1) **Bioset Criteria** passes Agilent QC flag and |log2 FC| > 1.5 and q-value < .05 ICL004-R VN1203 (H5N1), VN 7 hours, 1 MOI, Calu-3 cells Immport Host Factor ID Entrez Gene ID Public Identifier Bloset Information Key Log2 FC P-Value Log <u>Symbol</u> Name 100 2.2E-2 2.7 ABCA11P atp-binding EF. A 23 P92602 79963 NR 002451 2.0 cassette, sub-family a (abc1), member 11 (pseudogene) □ ABT1 activator of basal P A 23 P30784 29777 NM_013375 1 1.6 1.8E-2 1.9 transcription 1 ACRC acidic repeat EF. A 23 P171237 93953 NM 052957 1 2.2 1.3E-2 3.0 containing ACTA1 actin, alpha 1, P A 23 P1102 58 NM 001100 31.2 2.8E-3 79. skeletal muscle ACTA2 actin, alpha 2, RP. A 23 P150053 NM_001613 6.2 3.2E-2 5.7 smooth muscle, aorta ₽. 1.2E-2 9.5 ACTC1 actin, alpha, cardiac A 23 P205894 NM_005159 4.1 muscle 1 ACTG2 Ø. A 23 P39955 72 1 6.1 1.6E-2 49. actin, gamma 2, NM_001615 smooth muscle. enteric ACYP2 acylphosphatase 2, P. A 24 P336848 NM_138448 1 1.7 2.1E-2 2.8 muscle type ☐ AGAP1 图 116987 arfgap with gtpase A 32 P55161 NM 001037131 1 1.9 2.1E-2 2.7 domain, ankyrin repeat and ph domain 1 ☐ AGAP7 3 653268 2.0 5.9E-3 4.9 arfgap with gtpase A 24 P729905 NM 001077685 1 domain, ankyrin repeat and ph domain 7 E AGBL2 atp/gtp binding 4 A 32 P167705 79841 NM_024783 1 2.8 1.2E-2 3.6

protein-like 2

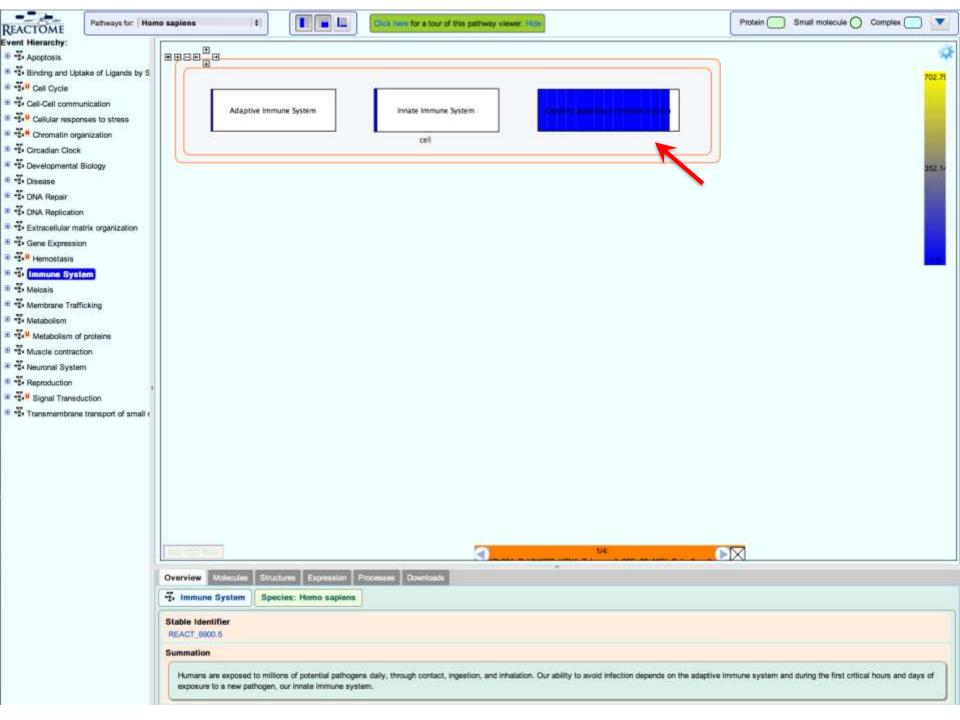


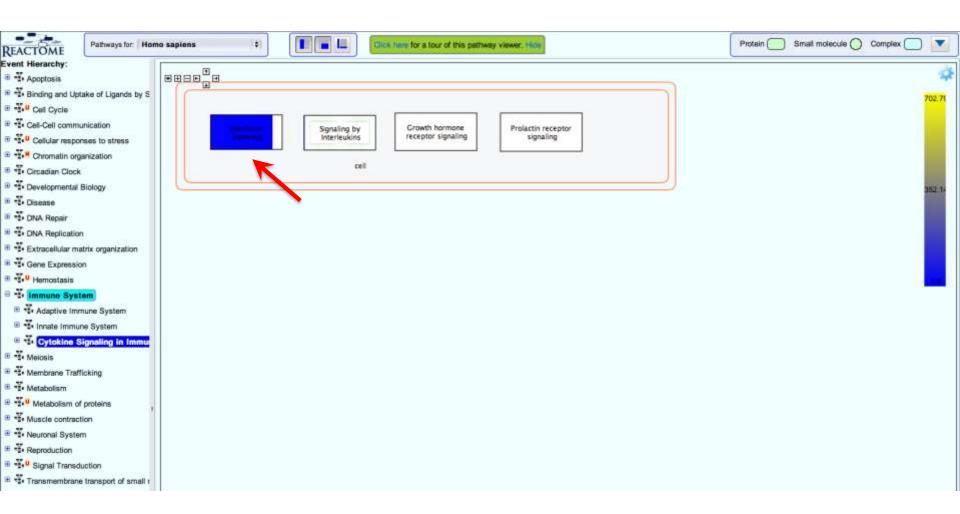
Analysis results, per pathway

sapiens

This table provides an overview of your expression data in a pathway context. For each Reactome pathway, the total number of proteins is shown, plus of the pathway, where your expression levels are represented as coloration of proteins.

Select format to do	ownload th	is table:	microsoft xcel (tsv) # Download	
Pathway **	Species	IDs in pathway (%)	CL004_R_VN1203_H5N1_7_hours_1_00E_00_MOI_Calu_3_cel	ls ICL004_R_VN1203_H5N1_12_hours_1_00E_00_M
Not assigned	Not known	(0%)	9.6	25.6
Apoptosis	Homo sapiens	1 (0%)	2.1	1.7
Binding and Uptake of Ligands by Scaveng	Homo sapiens	3 (1%)	12.3	28.8
Cell Cycle	Homo sapiens	9 (2%)	1.8	2.8
Cell-Cell communication	Homo sapiens	O (D%)		
Cellular responses to stress	Homo sapiens	1 (0%)	1.6	1.8
Chromatin organization	Homo sapiens	4.(496)	1.9	3.1
Circadian Clock	Homo sapiens	O.(D96)		
Developmental Biology	Homo sapiens	7 (1%)	2.8	4.5
Disease	Homo sapiens	23 (2%)	5.4	13.8
DNA Repair	Homo sapiens	1 (096)	1.6	2.8
DNA Replication	Homo sapiens	2 (2%)	1.9	3.3
Extracellular matrix organization	Homo sapiens	5 (296).	5.4	17.7
Gene	Homo	24 (3%)	2.4	4.2
Expression	sapiens	40.000	0.0	32.0
Hemostasis	Homo sapiens	16 (3%)	The state of the s	
Immune System	Homo	36 (3%)	9.6	22.0

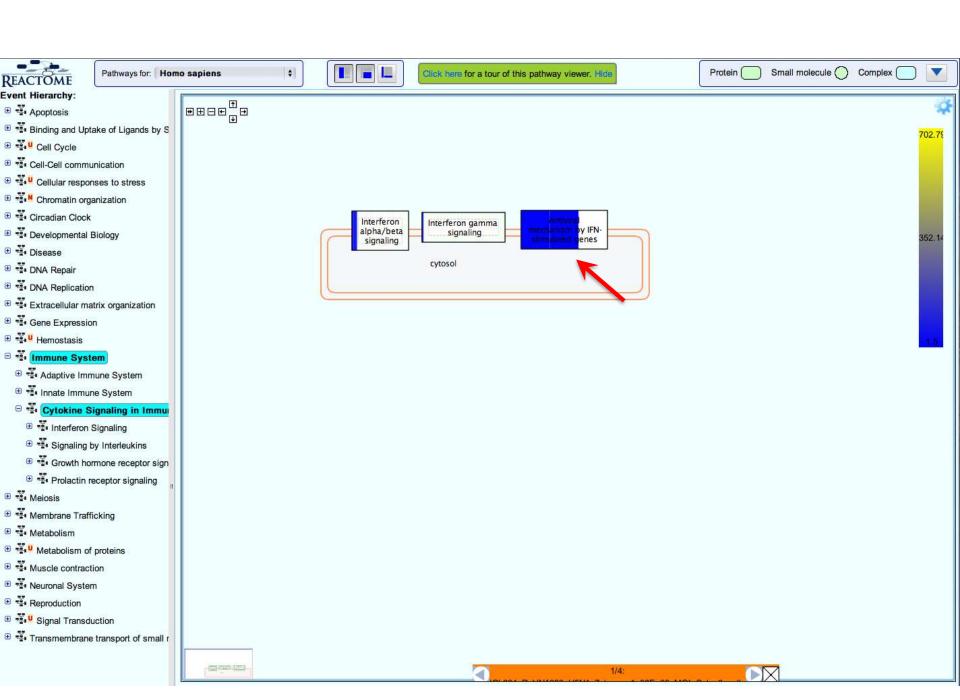


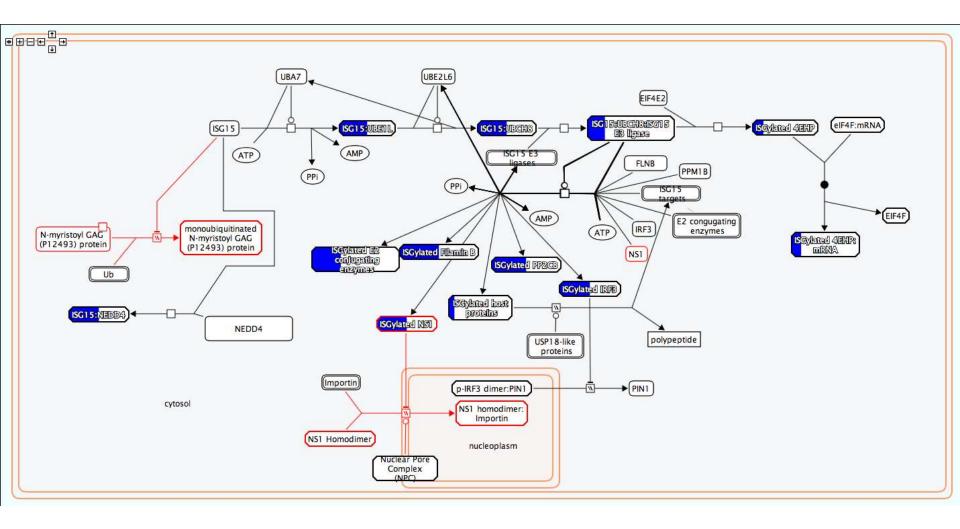


















Significant Host Factors as Compared to Control

"+" up-regulated, "-" down-regulated, blank: no change

Find expression pattern: RNAseL

SYMBOL

Find

Use comma to separate multiple entries. Ex. DDX58

Your search returned 55 records.

Data sorted by Number of Probes in descending

		order										
-	Pattern	Virus Time Post Infection Viral Dose Strain/Line	VN1203 (H5N1), 0 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 3 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 7 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 12 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 18 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 24 hours, 1 MOI, Calu-3 cells				
		Host Factors	0	0	<u>5277</u>	<u> </u>	<u>12944</u>	□ 17324				
	0,0,0,+,+,+	2921				+	+	+				
	0,0,0,-,-,-	2764				-	-	-				
	0,0,0,0,0,+	2105						+				
	0,0,0,0,0,-	1974						-				
	0,0,-,-,-,-	1643			-	-	-	-				
	0,0,0,0,-,-	1621					-	-				
	0,0,+,+,+,+	1094			+	+	+	+				
	0,0,0,0,+,+	931					+	+				
	0,0,0,+,0,0	919				+						
	0,0,-,0,0	746				-						
	0,0,0,-,0,-	487				-		-				
	+,0,+,0,0	452				+		+				
	0,0,+,0,0	433			+							
	0,0,0,+,+,0	433				+	+					
	0,0,0,-,-,0	402				-	-					
	0,0,-,0,0	275			-							
	0,0,0,0,-,0	230					-					
	0,0,-,-,0,0	222			-	-						
	0,0,-,-,-,0	195			-	-	-					
	+,0,0,+,0,0	192			+			+				
	0,0,-,-,0,-	180			-	-		-				
	0,0,0,0,+,0	152					+					
	0,0,+,0,+,+	133			+		+	+				
	0,0,0,+,0,-	115				+		-				
	0,0,+,+,0,0	110			+	+						
	0,0,-,0,-,-	102			-		-	-				
	0,0,-,0,0,-	102			-			-				

Significant Host Factors as Compared to Control

"+" up-regulated, "-" down-regulated, blank: no change

Find expression pattern:

SYMBOL RNAseL

Find

Use comma to separate multiple entries.

Ex. DDX58

Your search returned 1 record.

Data sorted by Number of Probes in descending

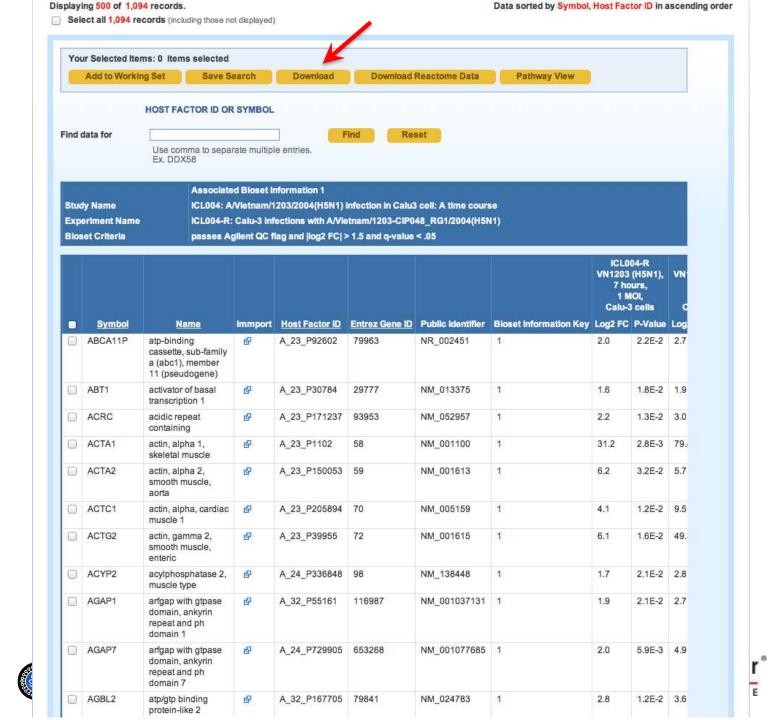
order

Pattern	Virus Time Post Infection Viral Dose Strain/Line	VN1203 (H5N1), 0 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 3 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 7 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 12 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 18 hours, 1 MOI, Calu-3 cells	VN1203 (H5N1), 24 hours, 1 MOI, Calu-3 cells	
	Host Factors	0	0	□5277	□ 13030	<u> </u>	□ 17324	
0,0,-,-	180			-	-		-	









3.	Experiment variable value 1	log2 Fold Change	Adjusted p- value	Experiment variable value 1	log2 Fold Change	Adjusted p- value	Experiment variable value 1	log2 Fold Change	Adjusted p- value	Experiment variable value 1	log2 Fold Change	Adjusted p- value	Experiment variable value 1	log2 Fold Change	Adjusted p- value
11813	13	-1.131153886	0.811597139	14	1.564843454	0.053754389	12	2.045130419	0.04537852	18	2.098974965	0.063960151	24	1.758797907	0.002917
11814		-1.004883272	0.987697131		-1.686246762	0.004212621	12	-1.760109848	0.009058148	18	-2.85109639	0.070304281	24	-6.338681279	0.020403
	Line .			170			12			18			24		
11815	Line -	-1.127237747	0.639828379		-1.038181834	0.877468486	12	-1.520819821	0.030117971	18	1.390211743	0.132987206	24	2.472705842	0.023203
11816		-1.132914897	0.804156099		1.110272302	0.869699135	12	2.408836912	0.037394034	18	2.469927467	0.251833771	24	5.483019801	0.018734
11817		1.113506963	0.485327841		-1.324509552	0.104280784	12	-1.914464291	0.012503302	18	-1.522732301	0.098559408	24	-1.769301005	0.033294
11818	3	-1.228789891	0.284346617	7	-1.498997532	0.062885106	12	-1.75233759	0.012899142	18	-1.516841556	0.091982819	24	-2.112064909	0.035343
11819	3	1.002137203	0.992041697	7	-1.102622134	0.662929877	12	-1.761783745	0.009011575	18	1.020812186	0.853205672	24	2.446585731	0.001137
11820	3	-1.168668591	0.537514164	7	1.668169818	0.225966254	12	-2.326971916	0.014047249	18	-2.643219438	0.316178618	24	-9.223953805	0.007485
11821	3	1.209027519	0.413739971	7	-1.819234161	0.041904688	12	-4.056013998	0.029811136	18	-8.550235918	0.075582847	24	-9.165251603	0.008939
11822	3	-1.097699367	0.618458938	7	1.531791165	0.14542335	12	1.638863123	0.019141701	18	1.092643552	0.806111263	24	-1.658466032	0.039107
11823	3	1.06914455	0.527936013	7	-1.312241892	0.072980309	12	-1.559867295	0.012568074	18	-1.069599722	0.858047926	24	-1.54511637	0.035452
11824	3	1.130611447	0.633790421	7	-1.900459964	0.029976993	12	-2.468589581	0.023690888	18	-12.59293969	0.053867539	24	-17.95333397	0.000419
11825	3	1.098535732	0.909402419	7	1.453425509	0.135134607	12	2.042578599	0.02670337	18	4.473426677	0.050539179	24	8.956037481	0.005123
11826	3	1.090593643	0.808766528	7	-2.555013274	0.010278205	12	-4.161120859	0.004358051	18	-6.203653108	0.112976889	24	-21.96429584	0.000899
11827	3	1.08654734	0.648306248	7	-1.533209381	0.063091572	12	-2.227653343	0.047516398	18	-10.81450906	0.067059062	24	-26.04841199	0.008638
11828	3	-1.617772682	0.325873477	7	-2.557499535	0.014259769	12	-7.185046268	0.023849116	18	-6.073874347	0.050403426	24	-26.92780274	0.006341
11829	3	-1.045038213	0.975752662	7	1.342812942	0.148028276	12	2.051808141	0.00262082	18	1.518148851	0.298996804	24	3.1993322	0.003295
11830	3	-1.599270411	0.349883277	7	-3.921198424	0.046013075	12	-2.431202577	0.004957721	18	-2.623164974	0.056005091	24	-4.711879967	0.032704
11831	3	1.021859105	0.962661537	7	1.259118526	0.189597341	12	1.749884772	0.023677065	18	2.174157429	0.125668602	24	4.485621874	0.000691
11832	3	-1.158952352	0.913298374	7	1.206971838	0.595342968	12	-2.371287468	0.030042671	18	-1.832819169	0.562190483	24	-3.4809188	0.013807
11833	3	-1.116677039	0.762525075	7	-1.314959666	0.119090741	12	-1.982368986	0.028192209	18	-2.802748916	0.053649918	24	-6.408424269	0.003045
11834	3	-1.094689556	0.959393828	7	-1.51784673	0.171560128	12	-1.509503638	0.037470011	18	-6.925109377	0.084255225	24	-18.60189989	0.004385







Summary of "Omics" Data Support in IRD/ViPR

- Structured metadata about study, experiments, analysis methods
- Series of derived biosets
- Boolean analysis of biosets from different experiments
- Biosets based on expression patterns
- Search for expression patterns of specific genes
- Access to complete data matrix
- Data linkout to pathway knowledgebase







Big Data to Knowledge

Volume + Variety = Value

Variety = Metadata

Data + Metadata + Interpretation = Knowledge







Acknowledgement

- Lynn Law, Richard Green U. Washington
- Peter Askovich Seattle Biomed
- Brian Aevermann, Brett Pickett, Doug Greer, Yun Zhang JCVI
- Entire Systems Biology Data Dissemination Working Group, especially Jeremy Zucker
- NIAID (Alison Yao and Valentina DiFrancesco)
- Entire ViPR/IRD development team at JCVI and Northrop Grumman
- NIAID/NIH N01AI40041





