

GENOMESPACE



(tools x connections) + recipes = frictionless science

Michael Reich

February 5, 2016

Interoperable, integrative computational genomics

Need

Insights through integrative studies of all data types

Challenges

Flood of high throughput data

- genomic sequence
- global mRNA expression profiles
- copy number and LOH
- epigenetic data
- protein level and modification status
- metabolite profiles

Proliferation of tools

- Databases, visualization, and analysis
- Difficulty of getting tools to work together
- Access, analyze, visualize each data type separately

GENOME SPACE

An online community to share computational tools

The screenshot shows the GenomeSpace website homepage. At the top, there's a navigation bar with links for "User Login | Admin Logout: michaelr", "Search", and "What is GenomeSpace? Tools Recipes Recipes (New) Documentation Developers Support About". Below the navigation is a large "GENOME SPACE" logo with the tagline "Frictionless connection of bioinformatics tools". There are "Register" and "User Login" buttons. A central image shows a network of interconnected nodes and a 3D visualization of data. Below this, a "STATUS" bar indicates "02.03.16 07:00AM" and "All systems are operating normally". A "Citing GenomeSpace" section provides citation information: "To cite your use of GenomeSpace, please reference Qu K, Garamszegi S, Wu F, et al. *Nature Methods*. 2016 Jan 18. doi: 10.1038/nmeth.3732.". The "WHAT'S NEW" section features news items: "Announcing the New GenomeSpace Recipe Resource" (Posted by Ted Liefeld on Tuesday, February 02, 2016 at 05:47PM), "New Format Converters in GenomeSpace" (Posted by Ted Liefeld on Friday, March 06, 2015 at 09:18AM), and "In response to user requests we have added the following three new format converters to the GenomeSpace ecosystem." A "See All Posts" link is at the bottom.

www.genomespace.org

- 12,000 registered users
- Published in Nature Methods: Ku et al, 2016

BRIEF COMMUNICATIONS

Integrative genomic analysis by interoperation of bioinformatics tools in GenomeSpace

Kun Qui^{1,12}, Sara Garamszegi^{2,12}, Felix Wu^{2,12}, Helga Thorvaldsdóttir², Ted Liefeld^{3,2}, Marco Ocana^{2,3}, Diego Borges-Rivera^{2,3}, Nathalia Pochet^{2,3}, James T Robinson^{2,3}, Barry Demchuk², Tim Hull³, Gil Ben-Artzi^{2,3}, Daniel Blankenberg², Galt P Barber², Brian T Lee², Robert M Kuhn², Anton Nekrutenko², Eran Segal², Trey Ideker², Michael Reich^{2,3}, Avi Regev^{2,10}, Howard Y Chang^{1,11} & Jill P Mesirov^{2,3}

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format converters, relieved scientists of the burden of identifying and scripting the conversions. The GenomeSpace Recipe Resource is a growing set of high-utility use cases that demonstrate how to leverage multiple tools and serve as quick guides to analysis tasks. The website serves as a knowledge base, newsstand and forum of contact and help for the community of users and tool developers.

Initially seeded by a consortium of biology research labs and development teams of six popular bioinformatics tools (Cytoscape¹², Galaxy³, GenePattern⁴, Genomics⁵, the Integrative Genomics Viewer (IGV)⁶ and the UCSC Table Browser⁷), GenomeSpace has since expanded to include many more sources. Our consortium labs provided biological projects and analytical needs to drive GenomeSpace design and development. For example, we recapitulated the steps and results of published analyses^{8,9} within GenomeSpace (Supplemental Figs. 1 and 2), dissecting and visualizing the gene regulatory networks in human cancer stem cell lines⁸ and the gene expression patterns in *S. cerevisiae*⁹. The study required diverse data types, analytical steps, and methods and multiple data transfers between tools. While originally requiring substantial scripting, this work can now be performed by non-programming biologists using only the GenomeSpace platform and tools within the browser.

The integrative analysis of diverse data types with multiple software tools remains an enormous challenge for many biologists. There is an ever-growing gap between the need to use various analysis and visualization tools and the complications of getting tools from different sources to work together. Moreover, it is difficult for experts, but especially for less experienced and computationally oriented biologists—to keep up with all of the available tools and to identify the right recipes to follow, particularly in the absence of an accepted “laboratory manual” for analytical protocols.

The integrative analysis of diverse data types with multiple software tools remains an enormous challenge for many biologists. There is an ever-growing gap between the need to use various analysis and visualization tools and the complications of getting tools from different sources to work together. Moreover, it is difficult for experts, but especially for less experienced and computationally oriented biologists—to keep up with all of the available tools and to identify the right recipes to follow, particularly in the absence of an accepted “laboratory manual” for analytical protocols.

GenomeSpace: Interoperable, Integrative Genomic Analysis



Lightweight “connection layer” between tools and data sources, with automatic format conversion.

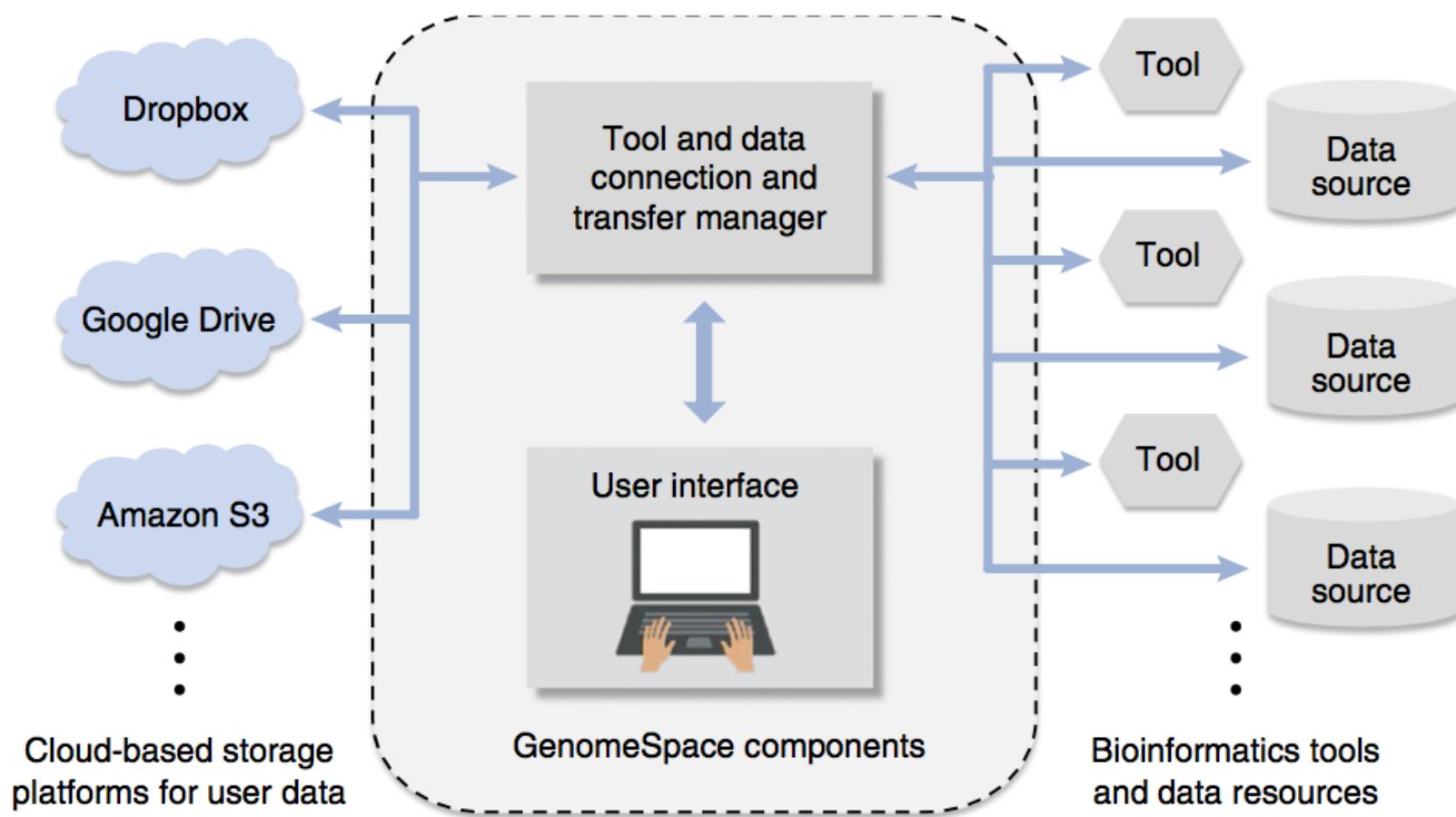


Cloud-based storage makes tools and data accessible from any location.

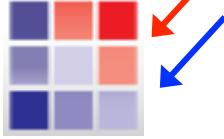
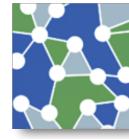
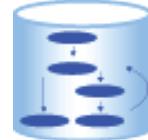


Tools retain their native look and feel.

GenomeSpace Architecture

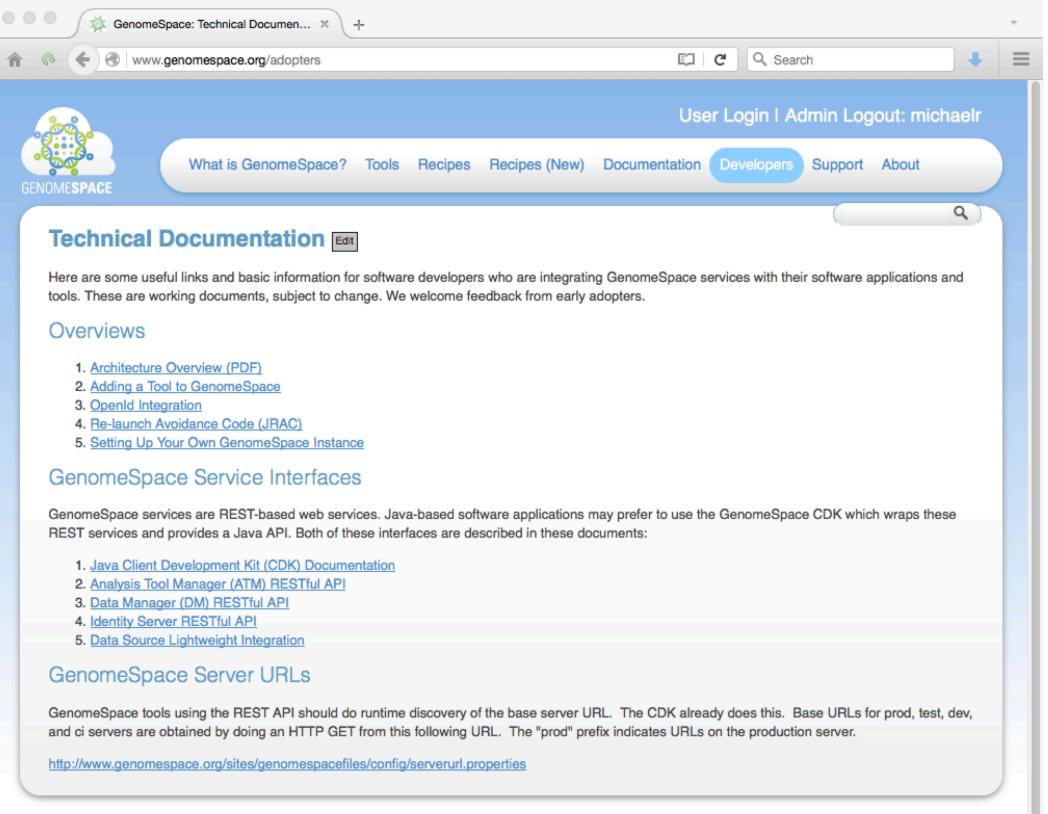


GenomeSpace Tools 2/16

				 Trinity
ArrayExpress	Cancer Cell Line Encyclopedia	Cistrome	Cytoscape	Cytoscape 3
				
Galaxy	GenePattern	Genomica	geWorkbench	Gitools
				
IGV	InSilicoDB	ISACreator	Multiple Myeloma Genomics Portal	MSigDB
				
Project Achilles	Reactome	Synapse	UCSC Table Browser	cBio Portal

GenomeSpace-enablement of new tools

- Basic enablement in ≤ 1 programmer-day
- RESTful API
- Client Development Kit (CDK)
- Data Transfer Utilities
- Documentation



The screenshot shows a web browser displaying the 'Technical Documentation' page of the GenomeSpace website. The URL in the address bar is www.genomespace.org/adopters. The page features a navigation bar with links for 'What is GenomeSpace?', 'Tools', 'Recipes', 'Documentation', 'Developers' (which is highlighted in blue), 'Support', and 'About'. A user login message 'User Login | Admin Logout: michaelr' is visible. The main content area is titled 'Technical Documentation' and includes a list of useful links for software developers:

- 1. Architecture Overview (PDF)
- 2. Adding a Tool to GenomeSpace
- 3. OpenID Integration
- 4. Re-launch Avoidance Code (JRAC)
- 5. Setting Up Your Own GenomeSpace Instance

Below this, there's a section titled 'GenomeSpace Service Interfaces' which describes REST-based web services and Java-based software applications. It lists five API documents:

- 1. Java Client Development Kit (CDK) Documentation
- 2. Analysis Tool Manager (ATM) RESTful API
- 3. Data Manager (DM) RESTful API
- 4. Identity Server RESTful API
- 5. Data Source Lightweight Integration

Finally, there's a section titled 'GenomeSpace Server URLs' with a note about runtime discovery and a link to the configuration file:

GenomeSpace tools using the REST API should do runtime discovery of the base server URL. The CDK already does this. Base URLs for prod, test, dev, and ci servers are obtained by doing an HTTP GET from this following URL. The "prod" prefix indicates URLs on the production server.

<http://www.genomespace.org/sites/genomespacefiles/config/serverurl.properties>

Add your own private tools

GENOME SPACE BETA

File | Launch | View | Connect | Manage

Cytoscape | ArrayExpress

Home

- ted
- Shared to ted
- dropbox:Ted Liefeld
- Public
- s3:1000genomes

Up to:

Edit HaploViewPlink

Name: HaploViewPlink

Description: HaploView with GenomeSpace extension

Tool Provider: Broad

Base URL: <http://vcancerportal05.broadinstitute.org:3003/Haplovie launcher/hapl>

Help URL:

File Parameter: Parameter name: plinkFi Required:

Allow multiple files: Multiple file Delimiter: ,

example call with dummy files
<http://vcancerportal05.broadinstitute.org:3003/Haplovie launcher/haplview.jnlp?plinkFiles=https%3A//gs.org/File1.txt,https%3A//gs.org/File2.txt>

Available formats: mitab, mut, ndb, nnf, odf, owl

Formats this tool accepts:

Icon: ./toolicon/haplovie.tif

Share with groups: GS-Developers

---Select a Group---

add

bench | Gitoools | Haplovie

Owner | Size | Last Modified

stem | stem | stem

GenomeSpace in action: Identification of transcription factor regulators

Cell

Resource

Densely Interconnected Transcriptional Circuits Control Cell States in Human Hematopoiesis

Noa Novershtern,^{1,2,3,11} Aravind Subramanian,^{1,11} Lee N. Lawton,⁴ Raymond H. Mak,¹ W. Nicholas Haining,⁵ Marie E. McConkey,⁶ Naomi Habib,³ Nir Yosef,¹ Cindy Y. Chang,^{1,6} Tal Shay,¹ Garrett M. Frampton,^{3,4} Adam C.B. Drake,^{2,7} Ilya Leskov,^{2,7} Björn Nilsson,^{1,8} Fred Preffer,⁸ David Dombrowski,⁹ John W. Evans,⁸ Ted Liefeld,¹ John S. Smutko,⁹ Jason Chin,¹⁰ Michael A.落¹¹, Richard A. Young,^{2,4} Todd R. Golub,^{1,5,10} Aviv Regev,^{1,2,10,12,*} and Benjamin L. Ebert^{1,5,6,12,*}

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⁷Koch Institute for Integrative Cancer Research, Massachusetts Institute of Technology, Cambridge, MA 02139

⁸Massachusetts General Hospital, Boston, MA 02114, USA

⁹Nugen Technologies, San Carlos, CA 94070, USA

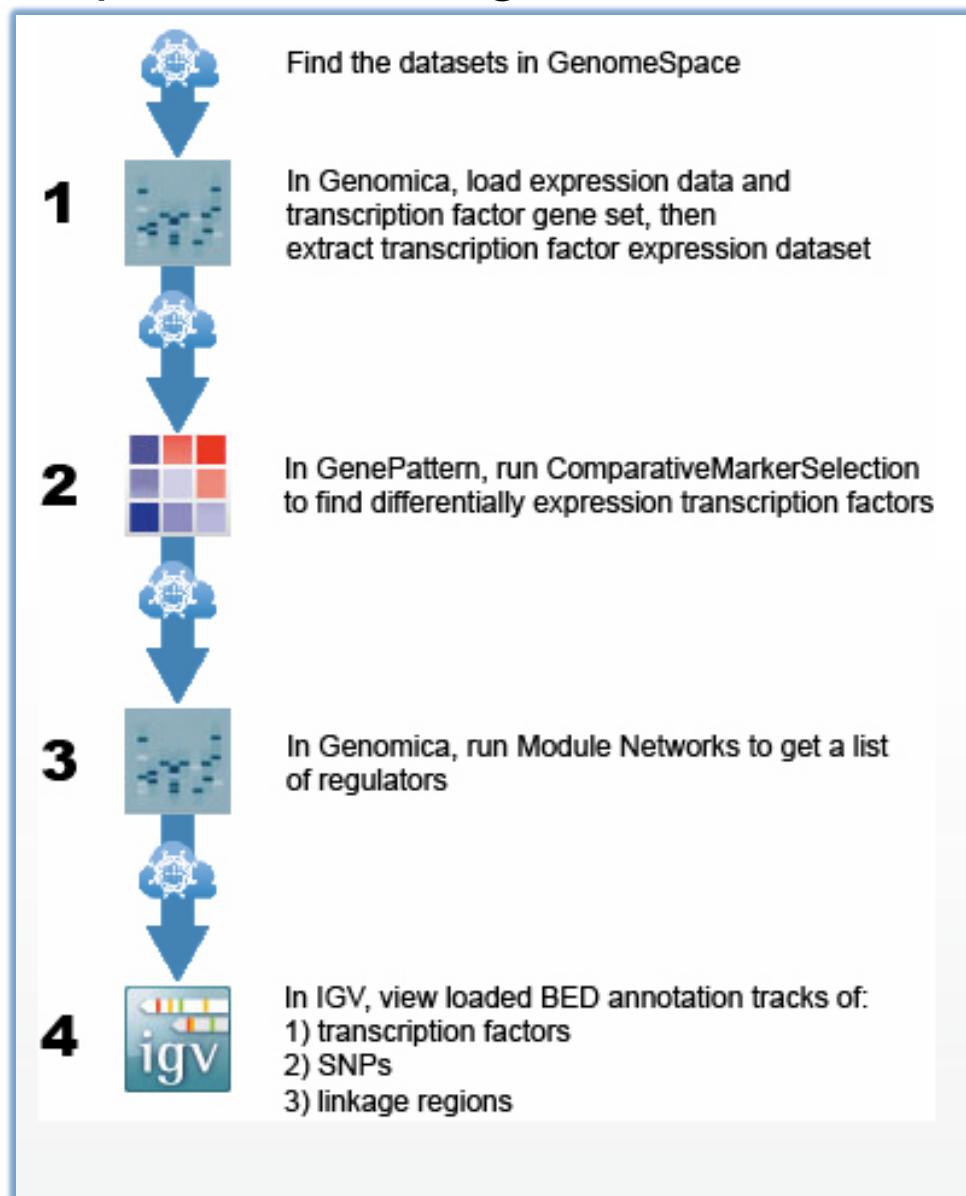
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¹¹These authors contributed equally to this work

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DOI 10.1016/j.cell.2011.01.004

Novershtern et al., Cell 2011

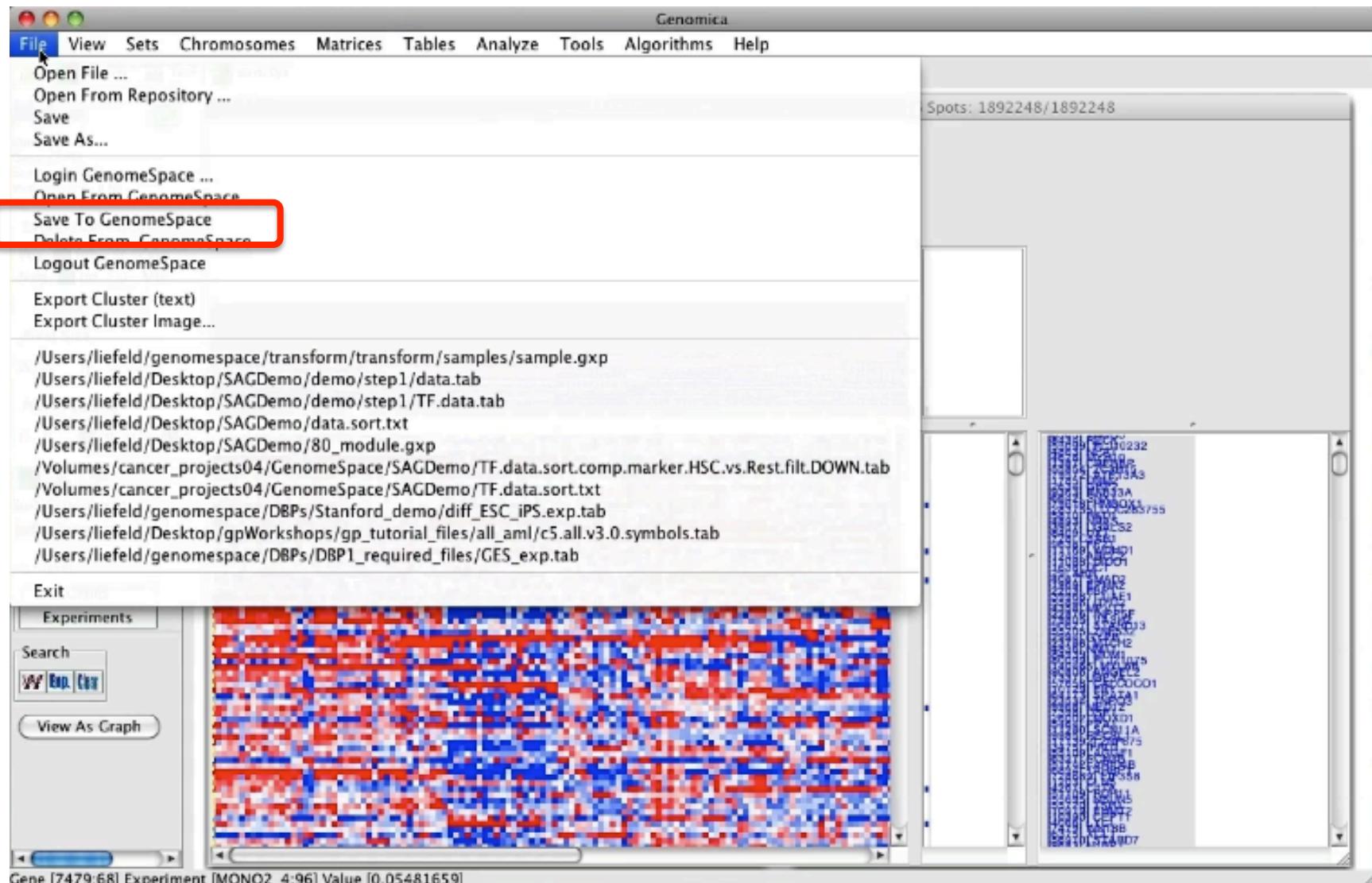


Doing this the standard way

1. Load file into Genomica
2. Run Genomica analysis to create transcription factor dataset
3. Download result file
4. Convert result file to GenePattern format
5. Upload file to GenePattern
6. Run GenePattern differential expression analysis
7. Download result file
8. Convert result file back to Genomica format
9. Upload to Genomica
10. Perform module network analysis
11. Download result file
12. Convert result file to IGV format
13. Open IGV
14. Load Genomica results into IGV
15. Visualize results

Script
Data transfer

Create transcription factor dataset in Genomica and save to GenomeSpace



Welcome to GenomeSpace

https://gsui.genomespace.org/gsui/gsui.html#

GENOME SPACE Invite a collaborator

File | Launch | View | Connect | Manage | Recipes | Help

Trinity GenePattern Genomica Cytoscape Reactome InSilicoDB Cytoscape 3 IGV ArrayExpress UCSC Table Browser Cistrome ISACreator Galaxy

Default

Home mmr

- + Analysis1
- + ArrayExpress
- + FGED
- + GSE10006GPL570_RNA_FRMAGENE_22280
- + GSE12790GPL570_RNA_FRMAGENE_2672
- + GSE8240GPL3921_RNA_ORIGINALGENE_17265
- + Illumina
- + MAGE-TAB
- + all_amr
- + all_amr_gsea
- + genomica_output
- + gs_workshop
- + gs_workshop2
- + shipp_lymphoma
- + step1
- step2**
- + step3
- + step4
- + Shared to mmr
- + dropbox:Michael Reich
- + Public

Up to: Home > mmr > **step2**

Filename	Tags	Owner	Size	Last Modified
DMAP.classes.cls		mmr	664 bytes	48 months ago
TF.data.comp.marker.HSC.vs.Rest.filt.gct		mmr	618 KB	48 months ago
TF.data.comp.marker.HSC.vs.Rest.filt.txt		mmr	1 KB	48 months ago
TF.data.comp.marker.HSC.vs.Rest.odf		mmr	126 KB	48 months ago
TF.data.gct		mmr	1.31 MB	48 months ago
TF.data.tab		mmr	1.31 MB	47 months ago
TF.data.txt		mmr	1.31 MB	48 months ago

Welcome to GenomeSpace

https://gsui.genomespace.org/gsui/gsui.html#

GENOME SPACE

Invite a collaborator

File | Launch | View | Connect | Manage | Recipes | Help

Trinity GenePattern Genomica Cytoscape Reactome InSilicoDB Cytoscape 3 igv ArrayExpress UCSC Table Browser Cistrome ISACreator Galaxy

Default

Up to: Home > mmr > step2

Filename	Tags	Owner	Size	Last Modified
<input checked="" type="checkbox"/> DMAP.classes.cls		mmr	664 bytes	48 months ago
<input type="checkbox"/> TF.data.comp.marker.HSC-vs.Rest.filt.gct		mmr	618 KB	48 months ago
<input type="checkbox"/> TF.data.comp.marker.HSC-vs.Rest.filt.txt		mmr	1 KB	48 months ago
<input type="checkbox"/> TF.data.comp.marker.HSC-vs.Rest.odf		mmr	126 KB	48 months ago
<input type="checkbox"/> TF.data.gct		mmr	1.31 MB	48 months ago
<input checked="" type="checkbox"/> TF.data.tab		mmr	1.31 MB	47 months ago
<input type="checkbox"/> TF.data.txt		mmr	1.31 MB	48 months ago

Home

mmr

- + Analysis1
- + ArrayExpress
- + FGED
- + GSE10006GPL570_RNA_FRMAGENE_22280
- + GSE12790GPL570_RNA_FRMAGENE_2672
- + GSE8240GPL3921_RNA_ORIGINALGENE_17265
- + Illumina
- + MAGE-TAB
- + all_amr
- + all_amr_gsea
- + genomica_output
- + gs_workshop
- + gs_workshop2
- + shipp_lymphoma
- + step1
- step2**
- + step3
- + step4

Shared to mmr

dropbox:Michael Reich

Public



Welcome to GenomeSpace

https://gsui.genomespace.org/gsui/gsui.html#

GENOME SPACE

Invite a collaborator

File | Launch | View | Connect | Manage | Recipes | Help

Trinity | GenePattern | Genomica | Cytoscape | Reactome | InSilicoDB | Cytoscape 3 | IGV | ArrayExpress | UCSC Table Browser | Cistrome | ISACreator | Galaxy

Default

Up to: Home > mmr > step2

Tags

Owner	Size	Last Modified
mmr	664 bytes	48 months ago
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mmr	1 KB	48 months ago
mmr	126 KB	48 months ago
mmr	1.31 MB	48 months ago
mmr	1.31 MB	47 months ago
mmr	1.31 MB	48 months ago

Filename

DMAP.classes.cls
TF.data.comp.marker.HSC.vs.Restfilt.gct
TF.data.comp.marker.gct
TF.data.gct
TF.data.tab
TF.data.txt

Launch GenePattern

Currently Selected Files

/mmr/step2/DMAP.classes.cls
/mmr/step2/TF.data.tab

Drop files here

TF.data.tab will be converted to an acceptable format.

Launch (button circled in red)

Close

Home

mmr

- + Analysis1
- + ArrayExpress
- + FGED
- + GSE10006GPL570_RNA_FRMAGENE_22280
- + GSE12790GPL570_RNA_FRMAGENE_2672
- + GSE8240GPL3921_RNA_ORIGINALGENE_17265
- + Illumina
- + MAGE-TAB
- + all_amr
- + all_amr_gsea
- + genomica_output
- + gs_workshop
- + gs_workshop2
- + shipp_lymphoma
- + step1
- + step2
- + step3
- + step4

Shared to mmr

dropbox:Michael Reich

Public

Perform differential expression analysis in GenePattern

Welcome to GenomeSpace Receive GenomeSpace File +

genepattern.broadinstitute.org/gp/pages/genomespace/receiveFile.jsf?files=https%3A%2F%2Fdm.genomespace.org%2Fdatamanager%2Ffile%2FHome%2Fmmr

Search mmrtest2

GenePattern

Modules & Pipelines Suites Job Results Resources Help GenomeSpace 0.0 KB / 30 GB

GenomeSpace has sent you a file

DMAP.classes.cls
Save a Copy: Upload Directory Submit

TF.data.tab Converted to gct
Save a Copy: Upload Directory Submit

Send to Module: ComparativeMarkerSelection Submit

The screenshot shows the GenePattern web application interface. At the top, there's a navigation bar with links for 'Modules & Pipelines', 'Suites', 'Job Results', 'Resources', 'Help', and 'GenomeSpace'. A user profile 'mmrtest2' is visible on the right. Below the navigation, a message says 'GenomeSpace has sent you a file'. It lists two files: 'DMAP.classes.cls' and 'TF.data.tab Converted to gct', each with a 'Save a Copy' dropdown and a 'Submit' button. A third section, 'Send to Module', contains the text 'ComparativeMarkerSelection' followed by a 'Submit' button, which is highlighted with a red border. The bottom of the page includes links for 'About GenePattern | Contact Us' and a copyright notice: '©2003-2015 Broad Institute, MIT'.

Perform differential expression analysis in GenePattern

The screenshot shows the GenePattern ComparativeMarkerSelection module interface. The top navigation bar includes links for Welcome to GenomeSpace, GenePattern - ComparativeMarkerS..., and a search bar. The main menu has options like Modules & Pipelines, Suites, Job Results, Resources, Help, and GenomeSpace. A user profile 'mmrtest2' is visible in the top right.

The main content area displays the ComparativeMarkerSelection module version 10. It has a brief description: "Identify differentially expressed genes that can discriminate between distinct classes of samples." Below this, there are several input fields:

- input file***: A dropdown menu showing a selected file: "https://dm.genomespace.org/datamanager/file/Home/mmr/step2/TF.data.tab?dataformat=http://www.genomespace.org/datamanager/dataformat/gct". A red box highlights this field.
- cls file***: A dropdown menu showing a selected file: "https://dm.genomespace.org/datamanager/file/Home/mmr/step2/DMAP.classes.cls". A red box highlights this field.
- confounding variable cls file**: A section with "Upload File..." and "Add Path or URL..." buttons, and a "Drag Files Here" area. A red box highlights this section.
- test direction***: A dropdown menu set to "2 Sided".
- test statistic***: A dropdown menu set to "T-Test".
- min std**: An empty text input field.

On the right side of the module interface, there are "Reset" and "Run" buttons, and a "Batch" checkbox. The entire module interface is also highlighted with a large red box.

Send differentially expressed genes to Genomica

Welcome to GenomeSpace

genepattern.broadinstitute.org/gp/pages/index.jsf

GenePattern

mmrtest2

0.0 KB / 30 GB

Modules & Pipelines Suites Job Results Resources Help GenomeSpace

Modules Jobs Files GenomeSpace Refresh

GENOMESPACE

GSE10006GPL570_RNA_FRMAGENE_222
80

GSE12790GPL570_RNA_FRMAGENE_267
2

GSE8240GPL3921_RNA_ORIGINALGENE
_17265

Illumina

known_gene

Irpprc.bed

MAGE-TAB

mm_track.bed

mRNA_orig.preprocessed.gct

Normals_Leu.preprocessed.comp.marker
.filt.gct

Normals_Leu.preprocessed.comp.marker
.filt.slice.gct

QC.sig_diffExp_genes.slice.txt

QC.sig_diffExp_genes.txt

SavedFromGenomica

shipp_lymphoma

step1

step2

DMAP.classes.cls

TF.data.comp.marker.HSC-vs.Rest.filt
.gct

TF.data.comp.marker.HSC-vs.Rest.filt
.txt

TF.data.comp.marker.HSC-vs.Rest.odf

TF.data.gct

TF.data.tab

TF.data.txt

step3

step4

TF.data.comp.marker.HSC-vs.Rest.odf

TF.data.gct

Public

Shared to mmr

TF.data.gct

Delete File
Permanently delete this file.

Save File
Save a copy of this file to your local computer.

Send to GenomeSpace Tool

Send to Galaxy
Send this file from GenePattern to Galaxy using GenomeSpace.

Send to Genomica (highlighted with red box)
Send this file from GenePattern to Genomica using GenomeSpace.

Send to Gitools
Send this file from GenePattern to Gitools using GenomeSpace.

Send to IGV
Send this file from GenePattern to IGV using GenomeSpace.

Send to Trinity
Send this file from GenePattern to Trinity using GenomeSpace.

Send to Module as tab

ChIPSeq.CreateHeatmap
Generates a heatmap based on the ChIP-Seq signal extracted from a BAM file, according to the... ChIP Seq

ChIPSeq.CreatePlots
Generates a scatter plot and correlation matrix based on the ChIP-Seq signal extracted from... ChIP Seq

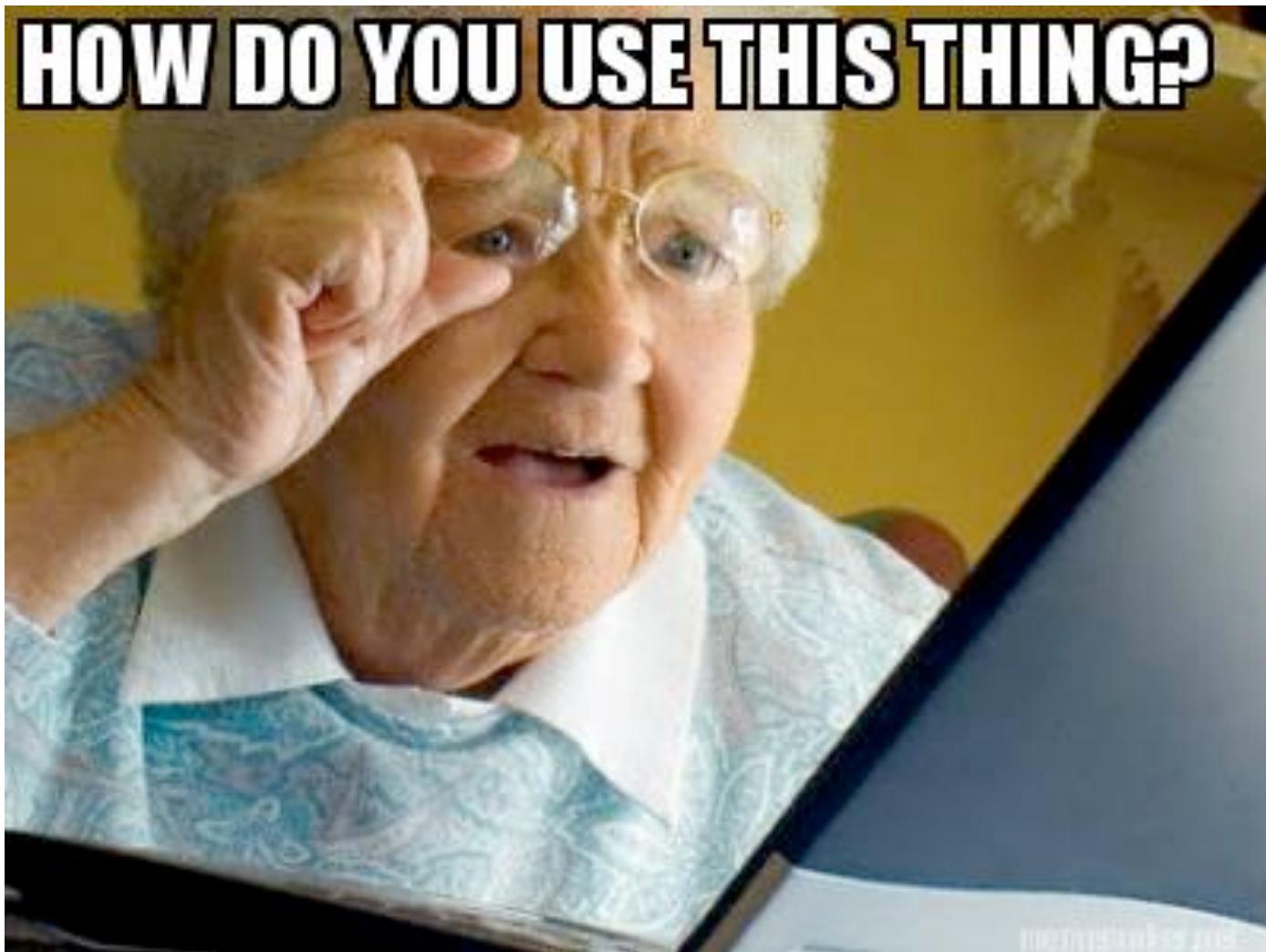
Search

0.0 KB / 30 GB

0.0 KB / 30 GB

GenePattern 2.0 Nature Genetics 38 no. 5 (2006): pp500-501 Google Scholar | Endnote | RIS

HOW DO YOU USE THIS THING?



GenomeSpace Recipe Resource

The image shows a screenshot of a web browser displaying the "Genomespace Recipe Resource" website at recipes.genomespace.org/home. The website features a "RECIPE RESOURCE" logo with a stylized globe icon. Below the logo, a brief introduction explains what GenomeSpace recipes are and how they can be used. A "Recipe Resource Guide" section provides links to "How to use the Recipe Resource and recipes", "Best Practices for Creating Recipes", and "Community Guidelines". A "Featured Recipe" section details a process for finding differentially expressed genes in RNA-Seq data, involving input (RNA-seq reads and reference gene annotations), analysis (aligning reads and identifying differentially expressed genes), and output (visualizing results with IGV). A large callout box highlights the use of the UCSC Table Browser and GenePattern. A detailed description of the recipe's purpose follows. At the bottom, a link invites users to "Click here to read the full recipe." To the right of the browser window, a portion of a book cover for "MOLECULAR CLONING: A LABORATORY MANUAL" by GREEN AND SAMBROOK is visible, specifically the fourth edition. The book cover features a grid of colorful circular icons.

GenomeSpace recipes provide step-by-step instructions for performing integrative bioinformatic data analyses using GenomeSpace tools and data sources. Recipes guide users through common analyses, starting with obtaining input followed by analyzing data by passing it to GenomeSpace-enabled tools. In addition, recipes guide users in interpreting the results of analyses.

To learn more about recipes, read the [Recipe Usage Guide](#) or try browsing recipes!

Recipe Resource Guide

- [How to use the Recipe Resource and recipes](#)
- [Best Practices for Creating Recipes](#)
- [Community Guidelines](#)

Featured Recipe

Find differentially expressed genes in RNA-Seq data

Which genes are differentially expressed between my two phenotypes based on my RNA-seq data?

Input: RNA-seq reads for multiple phenotypes, Reference gene annotations from the UCSC Table Browser

Analysis: Align the RNA-seq reads using reference gene annotations, identify differentially expressed genes, and process the aligned reads for visualization, using GenePattern

Output: Visualize aligned reads and differential expression using IGV

This recipe provides one method to identify differentially expressed genes in RNA-Seq read data. In particular, this recipe uses the UCSC Table Browser to retrieve a reference genome to align RNA-Seq reads against. We also use several modules in GenePattern to align the reads against the reference genome, and to identify differentially expressed genes when comparing two conditions. Finally, we use IGV to visualize the differentially expressed genes.

[Click here to read the full recipe.](#)

Browse Recipes

Search for GenomeSpace recipes

Contribute Recipes

[Read the Best Practices](#)

[Submit your recipe to the Recipe Resource](#)

[Check out our F1000 Genomics article](#)

Community

Do you have questions about other GenomeSpace resources?

Still have questions? [Contact us!](#)

FOCUS Research

FOURTH EDITION

MOLECULAR CLONING

A LABORATORY MANUAL

GREEN AND SAMBROOK

Current recipes

The screenshot shows the Genomespace Recipe Resource interface. At the top, there's a navigation bar with links for GenomeSpace Recipes, My Recipes, and GenomeSpace.org, along with a search bar and a login link. Below the header, a section titled "Expand all recipe descriptions" shows "Found 13 recipes". The recipes are listed in a grid format:

- Build and visualize a module network using putative aberrant regions and expression data** (Galaxy tool)
- Find differentially expressed genes in RNA-Seq data** (UCSC Table Browser and IGV tools)
- Find subnetworks of differentially expressed genes and identify associated biological functions** (Galaxy, UCSC Table Browser, and Cytoscape tools) **(highlighted with a red box)**
- Identify an up- or down-regulated pathway from expression data** (UCSC Table Browser, Galaxy, and MAGEOTools tools)
- Identify and validate a consensus signature using gene expression data** (UCSC Table Browser, Galaxy, and Cytoscape tools)
- Identify and validate coding variants from exome sequencing data** (Galaxy and IGV tools)
- Identify and visualize expressed transcripts in RNA-Seq data** (UCSC Table Browser, Galaxy, and IGV tools)
- Identify biological functions for genes in copy number variation regions** (Galaxy, Cytoscape, and MAGEOTools tools)
- Identify CNV-associated expression signatures using Stepwise Linkage Analysis of Microarray Signatures (SLAMS)** (Galaxy tool)

On the right side of the page, there are several filter panels:

- Filter by analysis type**: alignment, differential gene expression, functional annotation, gene expression analysis.
- Filter by data type**: copy number variation, DNA sequencing, gene sets, genomic loci.
- Filter by all available tags**: alignment, copy number variation, differential gene expression, DNA sequencing.
- Filter by tool**: ArrayExpress, CCLE, Cytoscape, Galaxy.

Recipe video and summary

Genomespace Recipe Resource +

recipes.genomespace.org/view/2

Find subnetworks of differentially expressed genes and identify associated biological functions

Added by sgaramsz on 2015.03.16

gene expression analysis network analysis differential gene expression microarray

0 0

Summary Inputs Overview Steps Results Interpretation Comments Back to the Top

1/9 Copying data from the Genome...

Summary

What subnetworks of differentially expressed genes are enriched in my samples? What biological functions are they related to?

Input Gene expression dataset Network annotations

Analysis Identify differentially expressed genes (DEGs) using *GenePattern* Build a network from the top DEGs using *Cytoscape*

Output Visualize subnetworks of highly interconnected DEGs

This recipe provides one method of using genes that are differentially expressed between two phenotypes, such as normal and tumor, to find subnetworks of interacting proteins and determine their functional annotations using Gene Ontology. In particular, this recipe makes use of several GenePattern modules to identify differentially regulated genes, then uses several Cytoscape plugins to identify potential interactions between gene products, and to visualize the resulting network.

Why differential expression analysis? We assume that most genes are not expressed all the time, but rather are expressed in specific tissues, stages of development, or under certain conditions. Genes which are expressed in one condition, such as cancer tissue, are said to be differentially expressed when compared to normal conditions. To identify which genes change in response to specific conditions (e.g. cancer), we must filter or process the dataset to remove genes which are not informative.

Why protein interaction network analysis? Gene expression analysis results in a list of differentially expressed genes, but it does not explain whether these genes are connected biologically in a pathway or network. To better understand the underlying biology that drives changes in gene expression analysis, we can perform network analysis to determine whether gene products (e.g. proteins) are reported to interact. To identify potential networks or pathways, we search for highly interconnected subnetworks within a large interaction network.

Inputs

To complete this recipe, we will need a gene expression dataset describing two conditions or phenotypes, such as cancer tissue vs. normal tissue. In this example, we will use gene expression data from a study in which committed granulocyte macrophage progenitor cells

Recipe steps

The screenshot shows a web browser window for the Genomespace Recipe Resource. The URL in the address bar is recipes.genomespace.org/view/2. The main content area displays a list of recipe steps, categorized by tool used: GenePattern and Cytoscape. Each step is numbered and described. At the bottom of the steps list are two buttons: '+ Expand All Steps' and '- Collapse All Steps'. To the right of the steps list is a sidebar with links to 'Summary', 'Inputs', 'Overview' (which is currently selected), 'Steps', 'Results Interpretation', 'Comments', and 'Back to the Top'.

Recipe steps

- **GenePattern**
 1. Loading data
 2. Filtering genes by expression value
 3. Identifying differentially expressed genes
 4. Selecting the top genes
 5. Extracting the top 50 genes
 6. Save the files to GenomeSpace
- **Cytoscape**
 7. Loading data into Cytoscape
 8. Identifying interacting proteins
 9. Finding differentially expressed subnetworks
 10. Exploring the subnetworks

+ Expand All Steps **- Collapse All Steps**

1: Loading data ➤
Load the files into GenePattern using one of the following methods.

2: Filtering genes by expression value ➤
We will use the [PreprocessDataset](#) module to filter out any genes that are not differentially expressed. In this recipe, we set the cut-off for differential expression at 3-fold up- or down-regulation. This module uses the GCT file.

3: Identifying differentially expressed genes ➤
We will use the [ComparativeMarkerSelection](#) module to identify genes which are differentially expressed and can distinguish between two phenotypes (e.g., normal vs. leukemic). This module uses the processed GCT file and the CLS file.

4: Selecting the top genes ➤
We will use the [ExtractComparativeMarkerResults](#) module to select the top genes that distinguish between phenotypes. In this recipe, we will extract the top 50 genes by rank.

5: Extracting the top 50 genes ➤
We will use the [SelectFileMatrix](#) module to select the gene names from our list of top 50 genes, allowing us to later import the file into Cytoscape. This module selects features from a file based on the rows and columns specified by the user. In this recipe, we will extract only the gene names.

6: Save the files to GenomeSpace ➤
Save the [Normal_Leu.genes.txt](#) file to GenomeSpace using one of the following methods.

7: Loading data into Cytoscape ➤
Load the data into Cytoscape.

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Single recipe step

Load the files into GenePattern using one of the following methods.

2: Filtering genes by expression value ▾

We will use the [PreprocessDataset](#) module to filter out any genes that are not differentially expressed. In this recipe, we set the cut-off for differential expression at 3-fold up- or down-regulation. This module uses the GCT file.

PreprocessDataset version 5

Performs several preprocessing steps on a res, gct, or odf input file

* required field

input filename* 2A 3

threshold and filter yes

Apply floor, ceiling and variation filter

floor 20

Value for floor threshold

ceiling 20000

Value for ceiling threshold

min fold change 3

Minimum fold change for variation filter

min delta 100

Minimum delta for variation filter

num outliers to exclude 0

Number of outliers per row to ignore when calculating row min and max for variation filter

row normalization no

Perform row normalization

1. Change to the [Modules](#) tab, and search for "PreprocessDataset".
2. Once the module is loaded, change the following parameters:
A. **input filename:** load the GCT file, e.g., [Normal_Leu.gct](#). To do this, navigate to the [GenomeSpace](#) tab, and navigating to the folder containing the GCT file. Load the file into the **input filename** parameter by clicking and dragging the file to the **input filename** input box.
3. Click **Run** to run [PreprocessDataset](#). This will generate a processed GCT file.

3: Identifying differentially expressed genes ▾

We will use the [ComparativeMarkerSelection](#) module to identify genes which are differentially expressed and can distinguish

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Create your own recipes

Genomespace Recipe Resource

recipes.genomespace.org/recipes/new

New GenomeSpace Recipe

Please fill out the form below to complete your recipe. Required fields are labeled as *****.

Title i *

Video URL (Please only use public video URLs) i

Recipe Title Image i
 No file selected.

Tags (comma delimited) i *

Summary i *

Inputs i *

GenomeSpace and ITCR

- Connecting more ITCR tools
- Demo projects using GenomeSpace
- GenomeSpace as a platform for interoperable informatics across ITCR



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