



# tranSMART

## User's Guide

**June 29, 2012**  
**Edition 1**



|                 | MRN       | DOB     | Age |   |   |         |
|-----------------|-----------|---------|-----|---|---|---------|
| Patient         | 100000001 | 8/13/71 | 37  | F | 1 | 8/9/10  |
| MRSA TEST, E-LA |           |         |     |   |   |         |
|                 | 100000001 | 7/23/79 | 33  | M | 1 | 7/29/08 |
|                 |           |         |     |   |   |         |
|                 |           |         |     |   |   |         |



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*Any blank pages in this document are intentionally inserted to allow correct double-sided printing.*

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# Getting Started with tranSMART

The tranSMART application reflects the efforts of various informatics groups to integrate data from internal and external data sources within a single data warehouse, and to provide scientific end users the tools to search for, view, and analyze the data in the warehouse.

The core internal data is a historical base of biomarker data from gene expression, RBM, and SNP experiments, including both raw and analyzed data.

External data sources include publicly available resources such as the Gene Expression Omnibus repository and MeSH Ontology.

The tranSMART application presents scientists with a search tool to query this vast ocean of disparate data through a Google-like user interface.

As users become more sophisticated in developing patterns of search terms, they can save and share their efforts with fellow researchers. A second tool, called the Dataset Explorer, allows the properly authorized user to create and study cohorts of patients that have been involved in completed clinical research efforts. Dataset Explorer also provides the user with tools to compare an individual (or group) in one study against a person or cohort in another study.



There may be some minor differences between the UI objects illustrated in this guide and the ones you see on your screen.

## Feature Overview

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tranSMART contains the following major features:

- Search tool
- Dataset Explorer
- Sample Explorer
- Gene Signature Wizard

## Search Tool

tranSMART provides a Google-like search tool that lets you search across multiple data sources for information related to items of interest, such as biomarkers, diseases, genes, and gene signatures.

The scope of a search can include clinical studies, externally conducted experiments, and in vivo/in vitro studies.

Search tool functionality includes:

- Searching within a particular category, such as diseases, genes, or pathways, or searching across all categories.
- Building complex search criteria that let you precisely define what to search for.
- Saving search criteria for easy recall and re-execution.
- Emailing search criteria to colleagues.

## Search Results

In searches of experiments, tranSMART displays complete listings of all analyses related to the experiments that are found.

tranSMART flags “meaningful” results in the analysis lists. Meaningful analyses are those where the signature genes are differentially modulated in a statistically significant way, indicating that the associated target is probably affected by the treatment, disease or other topic examined in the experiment.

Search result functionality includes:

- Displaying details of a particular experiment by clicking the name of the trial or experiment in the results list.
- Accessing a number of gene-related sites – Entrez Gene, Entrez Global, GeneCards, and Google Scholar – by clicking the name of a gene in the results list.
- Viewing the technical report or protocol used for an analysis.
- Exporting the complete results list to a Microsoft Excel file.
- Exporting details of a particular study, experiment, or other result to a Microsoft Excel file.

## Dataset Explorer

Dataset Explorer is an i2b2-based tool that lets you compare two sets of study groups based on one or more points of comparison. You define both the criteria that populate the study groups and the points of comparison between the study groups.

Dataset Explorer leverages the familiar navigation tree interface of Microsoft Windows Explorer to display data from clinical trials, and also leverages intuitive drag-and-drop functionality to help you build the criteria for populating the study groups and to add the points of comparison.

Dataset Explorer functionality includes:

- Saving the criteria used to populate the study groups.
- Emailing the study group criteria to colleagues.
- Using a heatmap to visualize the change in the expression of a specific protein from one sample to another.
- Using principal component analysis (PCA) to reduce the dimensionality of the dataset and to identify new, meaningful variables in the dataset.
- Performing advanced analyses and displaying results in various formats (scatter plot with linear regression, box plot with analysis of variance, etc.)
- Exporting a study or subset of a study to analyze in an external tool.

## Sample Explorer

Sample Explorer lets you search for datasets of tested tissue and blood samples, within categories such as tissue type, pathology, and test type (such as gene expression or SNP).

Once you find samples of interest, you can link back to the Dataset Explorer study for which the samples were collected.

## Gene Signature Wizard

tranSMART provides a wizard to help you create and define gene signatures and gene lists.

You can use your gene signature or gene list in tranSMART searches to find studies where the differentially regulated genes match those in your gene signature or list. This can help you develop hypotheses about diseases or treatments that may have similar genes deregulated.

Stored gene signatures can also be used in the analyses functionality of Dataset Explorer.

Gene signature functionality includes:

- Keeping the gene signature or list private so that only you can access it and use it in searches, or making it publicly available to all tranSMART users.
- Cloning an existing gene signature or list – either yours or a public one – as the starting point for creating and defining a new gene signature or list.
- Exporting all details of a gene signature or list to a Microsoft Excel file.

## Logging In

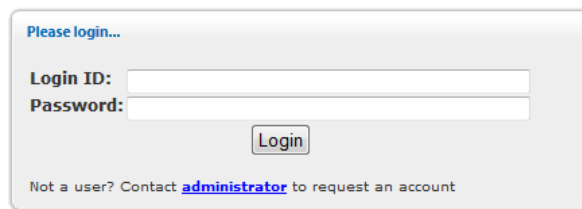
---

### To log into tranSMART:

1. Type the address of the tranSMART software into your browser's URL field:

<http://transmart.host.com/transmart>

The login screen appears:



2. Type your tranSMART login credentials, then click **Login**.

## Tools

---

tranSMART provides the following tools:

- **Search** – Search across internal and external data sources for research data and literature related to search terms that you provide.
- **Dataset Explorer** – View study data for subjects that you select, based on criteria that you specify. Also, compare data generated for subjects in two different study groups, based on criteria and points of comparison that you specify.
- **Sample Explorer** – Search for test samples using pre-defined search filters such as tissue, pathology, and dataset.
- **Gene Signature/Lists** – View definitions of existing gene signatures and add new gene signature definitions.



- **Utilities** – contains the following submenus:
  - **Help** – Display links to the tranSMART documentation set.
  - **Contact Us** – Email questions, problem reports, enhancement requests, or any other feedback about the tranSMART application.
  - **About** – Displays the version of tranSMART.

Select the tranSMART tool to use by clicking one of the tool tabs at the top of the tranSMART window:



## Opening a Particular Tool at Login

By default, tranSMART opens the Search tool after you log in. However, you can specify the tool for tranSMART to open immediately after login by including the tool name in the address you type into your browser's URL field.

To automatically open a particular tranSMART tool immediately after login, use an address listed below:



The addresses below are case-sensitive.

- Search tool – either of the following:
  - <https://transmart.host.com/transmart>
  - <https://transmart.host.com/transmart/search>
- Dataset Explorer tool
  - <https://transmart.host.com/transmart/datasetExplorer>
- Sample Explorer tool
  - <https://transmart.host.com/transmart/sampleExplorer>
- Gene Signature/Lists tool
  - <https://transmart.host.com/transmart/geneSignature>



## Chapter 2

# Search Tool

tranSMART provides a Google-like interface for searching across internal data sources as well as external data sources with a single query, based on one or more search filters that you define.

### Search Tasks

---

A search filter is the name of a biomedical concept such as a gene, pathway, disease, or other item of medical interest. These filter names are pre-defined in tranSMART. You can browse lists of these filter names and select the filter you want, or type part of a filter name in the **Search** field, causing tranSMART to display a list of filters that begin with the text you type.

You can base your search on a single search filter or on a multi-filter search string.

### Defining a Search Filter

There are several ways to define a search filter:

- Type all or part of the filter name directly into the **Search** field.
- Browse all the pre-defined filters within filter categories (such as diseases).
- Use a saved search filter or search string.

## Type the Filter Name

**To search the internal and external data sources for information related to a filter name:**

1. Click the tab for the Search tool at the top of the tranSMART window.
2. Click the search filter category to search within (for example, search only diseases, or search only genes).

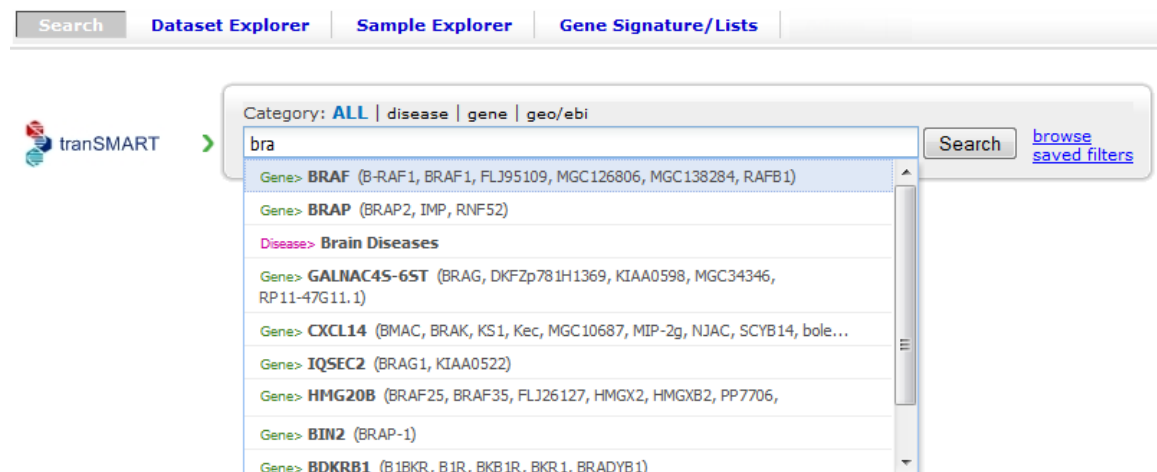
The search engine first filters by the filter category you select, and then filters by the name you type. To search across all filter categories, click **all**:



You can only specify one search filter in the **Search** field shown above. For instructions on creating a multi-filter search string, see [Building a Search String](#) on page 12.

3. Type part or all of the filter name into the **Search** field.

Up to 20 matches that begin with the text you type are displayed in a dropdown list below the **Search** field. For example, the following list appears for the search filter **bra** when searching across all filter categories:



You can also search for aliases. For example, to find the gene PTK7, you can type part or all of the name PTK7 or its alias, CCK4.

4. Do one of the following:
  - If the name of the filter you want appears in the list, click the filter name. The search begins immediately.
  - If the filter name you want does not appear in the list, type a more complete name in the **Search** field. For example, if you typed only **br** in the **Search** field, no entries for “brain diseases” appear in the list. Typing an **a** after the text you already typed displays a list like the one shown above.
  - If no list appears after you type a complete filter name, click the **Search** button.
5. To start another search using a new search filter, click **clear all** above the search result:

About 2 results found

Filters: Disease> Brain Diseases ✕ [advanced](#) [save](#) [clear all](#)

Alternatively, you can click the tranSMART logo, or simply type a new filter in the **Search** field.

See [Working with Search Results](#) on page 17 for information on viewing and refining search results.

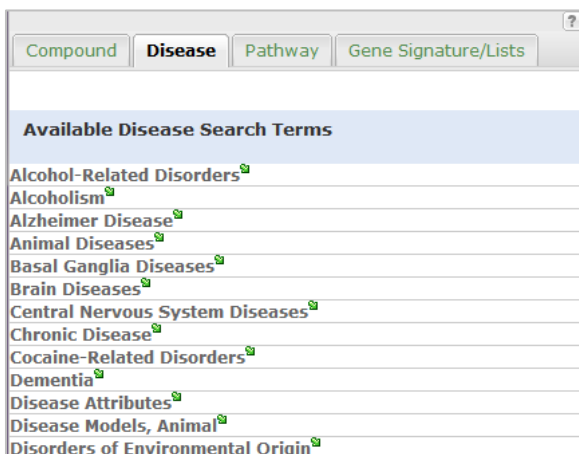
## Browse for a Filter Name

You can browse through all the pre-defined filters in each of the following areas:

- Disease
- Gene Signature/Lists
- Geo/ebi
- Pathway

### To browse the pre-defined filters:

1. Click the tab for the Search tool at the top of the tranSMART window.
2. Click the **browse** link to the right of the **Search** button. A window similar to the following appears:



The search engine ignores any filter category you may have selected and any filter text you may have entered in the **Search** field.

3. Click the tab for the area in which you want to browse for filters.
4. To initiate a search for information related to a filter, click the filter name or the green arrow (→) after the name.

After you click a filter, the search begins immediately.

5. To browse for another filter, click **browse** again. There is no need to clear the previous result.

- To start another search using a new search filter, click **clear all** above the search result:

About 2 results found

Filters: Disease> Brain Diseases  [advanced](#) [save](#) [clear all](#)

Alternatively, you can click the tranSMART logo, or simply type a new filter in the **Search** field.

See [Working with Search Results](#) on page 17 for information on viewing and refining search results.

## Use a Saved Search Filter

There are two ways to access a saved search filter:

- Retrieve the saved filter from a list of filters that you created and saved. The instructions in this section describe this method.
- Click a link to a saved filter that someone else has created, saved, and emailed to you.

See [Saving a Search Filter or Search String](#) on page 15 for more information, including instructions on saving search filters and search strings.

### To search against a filter that you created and saved:



- Click the tab for the Search tool at the top of the tranSMART window.
- Click the **saved filters** link to the right of the **Search** button. A list of filters that you created and saved appears:

#### Search > Saved Filters

##### demo filter

[select](#) [edit](#) [delete](#)

November 28th, 2011: I was looking for either TLR3 or IL4 and an endocrine sys disease

Genes> IL4  OR TLR3  AND Disease> Endocrine System Diseases

Shortcut: Private

- To search against a saved filter in the list, click the **select** link to the right of the saved filter name. The search begins immediately.
- To start another search using a new search filter, click **clear all** above the search result.

Alternatively, you can click the tranSMART logo, or simply type a new filter in the **Search** field.

See [Working with Search Results](#) on page 17 for information on viewing and refining search results.

## Building a Search String

You can make the scope of your search more precise by building a multi-filter search string. The filters in a search string are joined by the logical operators **AND** and **OR**.

### Rules for Building a Search String

The following rules apply to building a multi-filter search string:

- Filters within the same filter category (such as diseases or genes) are joined by the logical operator **OR**.

For example, if you add the filters **Diseases> Melanoma** and **Diseases> Melanoma, Experimental** to a search string, the search engine evaluates them as in the following expression:

```
(Diseases> Melanoma OR Diseases> Melanoma, Experimental)
```

- Filters within different filter categories are joined by the logical operator **AND**.

For example, if you add the filters **Diseases> Anemia**, **Diseases> Anemia, Hemolytic**, and **Gene> HBB** to a search string, the search engine evaluates them as in the following expression:

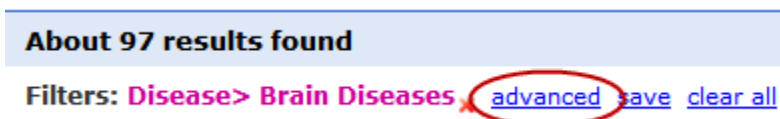
```
(Diseases> Anemia OR Diseases> Anemia, Hemolytic) AND Gene> HBB
```

- Filters that are not among the pre-defined filters are assigned to the filter category **Text>**.

### Instructions for Building a Search String

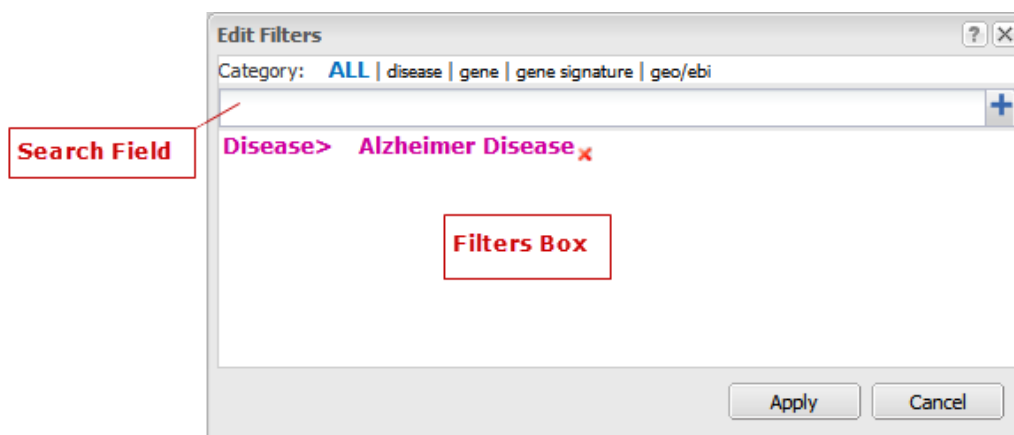
**To build a multi-filter search string:**

- Define a search filter using any of the methods described in [Defining a Search Filter](#) on page 7.
- When the results window appears, click **advanced**:



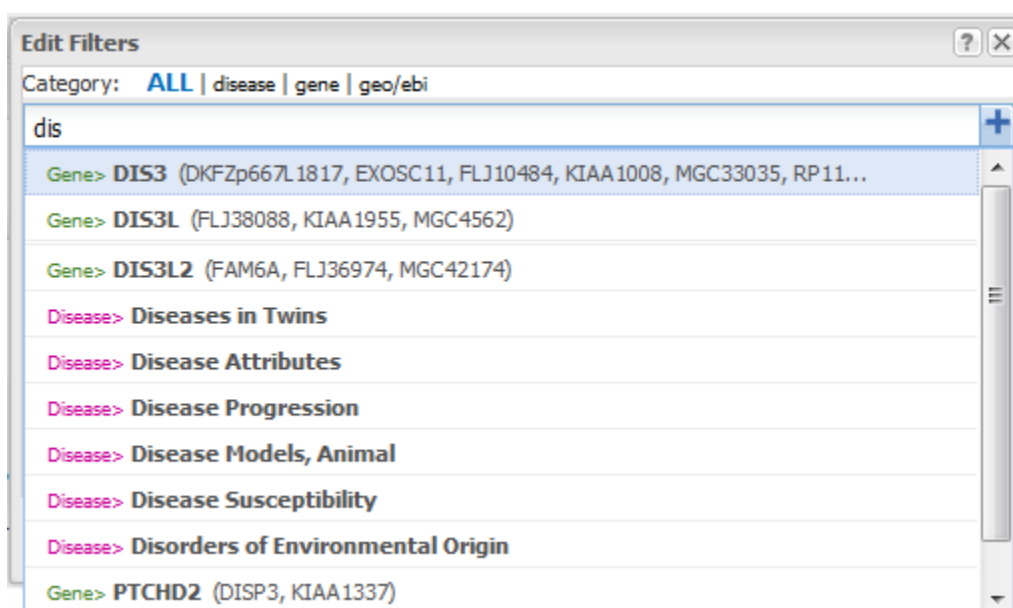


The Edit Filters dialog appears, displaying the filter you just created:




- To add another filter, type part or all of a filter name into the **Search** field.

Up to 20 matches for the text you type are displayed in a dropdown list below the **Search** field. For example, the following list appears for the search filter **dis**:

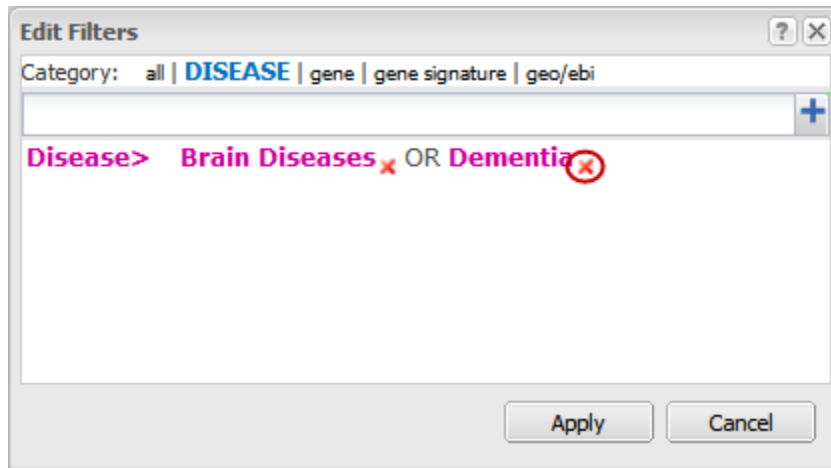


Do one of the following:

- ☐ If the name of the filter you want appears in the list, click the filter name. The tranSMART software inserts the filter into the **Filters Box**.
- ☐ If the filter you want does not appear in the list, type a more complete name in the **Search** field.
- ☐ If no list appears after you type a complete filter name, click the plus-sign button (  ) to the right of the **Search** field.

- Repeat the previous step for each new filter to add to the search string.

5. Optionally, to delete a filter from the search string, click the red **X** (✖) to the right of the filter name:



6. When finished defining the search string, click **Apply** to begin the search.
7. When the results window appears, you can continue editing the search string or save it, as follows:
  - To continue editing the search string, click **advanced**.
  - To save the search string, click **save**.

About 2 results found

Filters: Disease> Brain Diseases ✖ OR Dementia ✖ [advanced](#) [save](#) [clear all](#)

Continue editing  
the search string

Save the  
search string

The search engine evaluates this search string as in the following expression:

(Disease> Brain Diseases OR Dementia)

See [Saving a Search Filter or Search String](#) on page 15 for more information about saving search filters and search strings.

## Saving a Search Filter or Search String

### To save a search filter or search string:

1. After defining the search filter or search string, run the search and click **save** in the results window.

The Create Filter window appears:



**Create Filter**

Name:

Description:

Private Flag: ☐

Summary: Diseases> Brain Diseases OR Diseases in Twins

 **Create**  **Cancel**

2. In the **Name** field, type a name for the search filter or search string.
3. Optionally, in the **Description** field, type a description of the search filter or search string. In the saved filters list, the description appears immediately below the name of the search filter or search string.
4. Check the **Private Flag** checkbox to prevent others from using this search filter or search string, or clear the checkbox to allow others to use the search filter or search string.


If a filter is public, a shortcut (link) to the filter is displayed in the **saved filters** list, and an **email** link is provided, allowing you to email the shortcut to others. If a filter is private, the saved filter is marked "Private," and the filter shortcut and **email** link are not displayed.



Only the person who created and saved a search filter can see that filter in the saved filter list. To let a colleague use a search filter you saved, you must (1) mark the filter as Public, and (2) click the **email** link to send the shortcut for the search filter to the colleague.

In the following **Saved Filters** list, the first two entries are private and the third is public:

**Search > Saved Filters**

|  |
|--|
| KEGG Pathway and Melanoma  |
| Pathway> KEGG-Melanoma  AND Diseases> Melanoma OR Melanoma, Experimental AND Text> "Melanoma" |
| Shortcut: Private  |
| Melanoma   |
| Diseases> Melanoma OR Melanoma, Experimental AND Text> "Melanoma"  |
| Shortcut: Private  |
| Skin Cancer  |
| Excluding basal cell.  |
| Diseases> Carcinoma, Squamous Cell OR Melanoma AND Text> "Skin Cancer"   |
| Shortcut: /transmart/search/searchCustomFilter/1690018 <a href="#">email</a>   |

5. When finished, click **Create** to save the new search filter or search string, or click **Cancel** to abandon it.

## Editing and Deleting Saved Filters

### To edit a saved filter:

1. Click the tab for the Search tool at the top of the tranSMART window.
2. Click the **saved filters** link to the right of the **Search** button. A list of your saved search filters appears.
3. Click the **edit** link to the right of the saved filter name. The Edit Filter window appears:

**Edit Filter**

Name:

Description:

Private Flag: ☐

Summary: **Diseases> Carcinoma, Squamous Cell OR Melanoma AND Text> "Skin Cancer"**

Update Delete Cancel

4. Make one or more of the following changes:
  - ☐ In the **Name** field, modify the name of the saved filter.
  - ☐ In the **Description** field, add or modify an optional description of the saved filter. In the **saved filters** list, the description appears immediately below the saved filter name.
  - ☐ Check the **Private Flag** checkbox to prevent others from using this saved filter, or clear the checkbox to allow others to use the saved filter.

Another user can use a filter you created and saved only (1) if the filter is public, and (2) you email the user the shortcut (link) to the filter.

  - ☐ To delete the filter you are editing, click the **Delete** button ( ).



These are the only changes you can make to a saved filter. To make changes to the filter itself, run the search against the filter, then click **advanced** to define a new search filter based on the existing one. For details, see Instructions for [Building a Search String](#) on page 12.

5. When finished making changes, click the **Update** button to save your changes, or click the **Cancel** button to abandon them.

### To delete a saved filter from the saved filters list:

1. Click the tab for the Search tool at the top of the tranSMART window.
2. Click the **saved filters** link to the right of the **Search** button. A list of saved search filters appears.
3. Click the **delete** link to the right of the saved filter name.

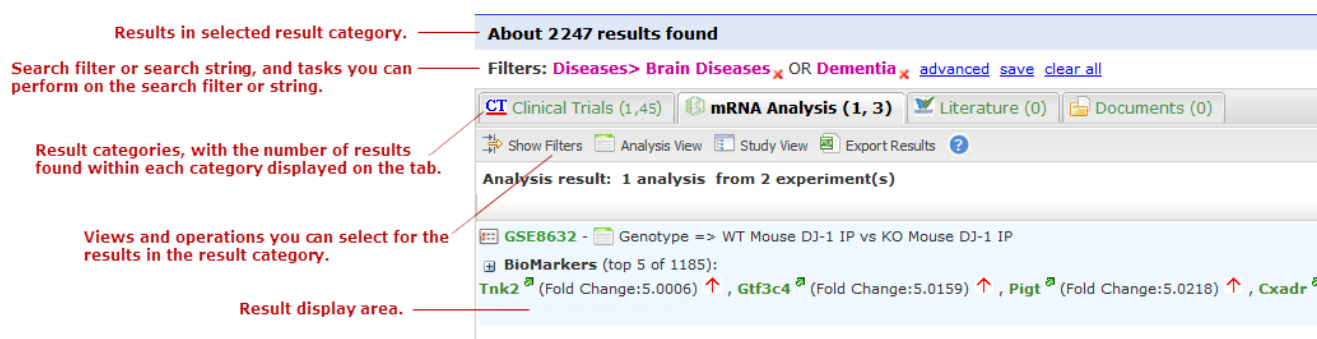
## Working with Search Results

The results window displays all the clinical, documentary, and other information found in the data warehouse that relates to the search filter or search string.

The content of the results window varies, depending on the result category you select (for example, Clinical Trials or mRNA Analysis) and the type of view you want to use to display the results (for example, Heatmap or Study View). Some result categories also let you further refine the results by adding more filters to the search.

To select a result category to view, click the tab that contains the result category name.

The following figure shows the sections of the results window:



The tabs for the result categories Clinical Trials and mRNA Analysis display pairs of numbers. These numbers represent the following results:

### ■ Clinical Trials ( x, y )

- x = the number of statistically significant analyses. These hits can be viewed in the Analysis View.
- y = the total number of analyses. These hits can be viewed in the Study View.

For example, in the preceding figure, 1 statistically significant analysis was returned, and a total of 45 analyses were returned.

### ■ mRNA Analysis (x, y)

- x = the number of statistically significant analyses. These hits can be viewed in the Analysis View.
- y = the total number of analyses. These hits can be viewed in the Study View.

For example, in the preceding figure, 1 statistically significant analysis was returned, and a total of 3 analyses were returned.

A *statistically significant analysis* is one in which the genes in a gene signature, gene list, or pathway are differentially modulated in a statistically significant way, indicating that the associated target or pathway is probably affected by the treatment, disease or other topic examined in the study.

To qualify as a statistically significant analysis, certain data points (such as p-value) must be evaluated and attain an aggregate score that meets or exceeds a particular

threshold, based on an internal algorithm. For information on the rules that determine how analysis results are ranked, see [TEA Analyses](#) on page 24.

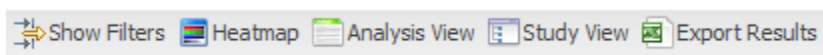
The following sections describe the views and operations available for each result category:

- [Clinical Trials Tab](#) (Page 18)
- [mRNA Analysis Tab](#) (page 19)
- [Literature Tab](#) (page 24)
- [Documents Tab](#) (page 24)

## Clinical Trials Tab

This result category contains data from internal clinical trials.

Click the **Clinical Trials** tab to display the results in this category. The buttons in the following figure appear at the top of the results list. You may see fewer buttons, depending on the results of your particular search:



These buttons give you access to the following views and operations:

- **Show Filters** – Define additional filters to further refine the search results.
- **Heatmap** – View the results as a heat map.
- **Analysis View** – View a list of the statistically significant analyses of the clinical trials.

Results are sorted from the highest-scoring analysis down to the lowest.

- **Study View** – View a list of the clinical trials and, optionally, *all* the analyses for each clinical trial – that is, those analyses that are considered statistically significant and those that are not.

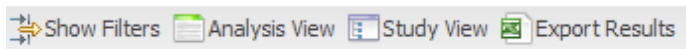
Results are sorted from the clinical trial with the most matches with the search filter or search string, down to the one with the least matches.

- **Export Results** – Export descriptions of each clinical trial, and also all the analysis data from each of the clinical trials, to a Microsoft Excel file. All clinical trial descriptions are written to one worksheet in the file, and all analysis data is written to a second worksheet in the file.

## mRNA Analysis Tab

The mRNA Analysis tab contains gene expression data derived largely from external experiments and from some internal experiments.

Click the **mRNA Analysis** tab to display the results in this category. The buttons in the following figure appear at the top of the results list. You may see fewer buttons, depending on the results of your particular search:



These buttons give you access to the following views and operations:

- **Show Filters** – Define additional filters to further refine the search results.
- **Analysis View** – View the analyses of the experiments that are ranked as statistically significant analyses.
- **Study View** – View the details of the experiments and, optionally, *all* the analyses for each experiment – that is, those analyses that are considered statistically significant and those that are not.
- **Export Results** – Export descriptions of each experiment, and also all the analysis data from each of the experiments, to a Microsoft Excel file. All descriptions of experiments are written to one worksheet in the file, and all analysis data is written to a second worksheet in the file.

The following sections describe the results of experiments for the disease **Amyotrophic Lateral Sclerosis**. Click the **mRNA Analysis** tab to see the results.

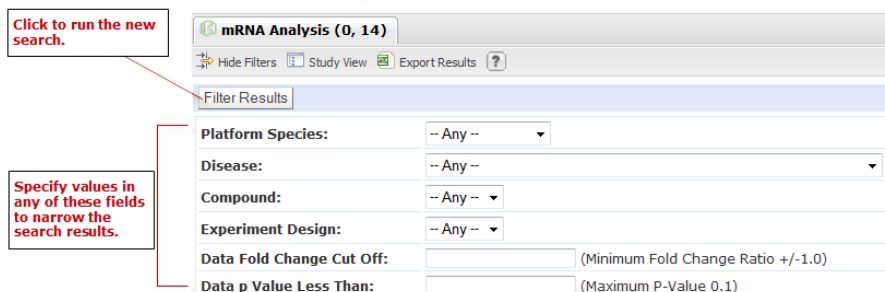
### Show Filters

Click the **Show Filters** button to further refine the search results. When you click the button, a window containing filter fields appears (shown below), and the **Show Filters** button is replaced by the **Hide Filters** button.

In the figure below, filter selections are set for the broadest possible search.

### To narrow the search:

1. Specify one or more filters – for example, specify a particular p-value to search against, and/or select a particular disease from the dropdown list.
2. Click **Filter Results** to start the search.



## Analysis View

Click the **Analysis View** button to view the statistically significant analyses associated with mRNA experiments.

For information on the rules that determine how analysis results are ranked, see [TEA Analyses](#) on page 24.

Experiment. Click or hover cursor to open details box.

Description of experiment. Click or hover cursor to open details box.

About 4 results found

Filters: Disease> Amyotrophic Lateral Sclerosis [advanced](#) [save](#) [clear all](#)

mRNA Analysis (5, 6)

Show Filters Analysis View Study View Export Results

Analysis result: 5 analyses from 4 experiment(s)

GSE833 - DiseaseState => sporadic ALS vs control

BioMarkers (top 5 of 118):

CAPG (Fold Change:221.8659) , FOXO4 (Fold Change:215.6599) , CD14 (Fold Change:193.2086) , PRKCSH (Fold Change:190.8862) , PRKCD (Fold Change:134.5789)

| CAPG   |  | ProbeSet: M94345_at      | Gene: CAPG   | Fold Change: 221.866 | p-Value: 0 |
|--------|--|--------------------------|--------------|----------------------|------------|
| FOXO4  |  | ProbeSet: X93996_rna1_at | Gene: FOXO4  | Fold Change: 215.66  | p-Value: 0 |
| CD14   |  | ProbeSet: X13334_at      | Gene: CD14   | Fold Change: 193.209 | p-Value: 0 |
| PRKCSH |  | ProbeSet: U50327_s_at    | Gene: PRKCSH | Fold Change: 190.886 | p-Value: 0 |
| PRKCD  |  | ProbeSet: D10495_at      | Gene: PRKCD  | Fold Change: 134.579 | p-Value: 0 |

GSE833 - DiseaseState => familial ALS vs control

BioMarkers (top 5 of 104):

CAPG (Fold Change:858.1024) , AQP1 (Fold Change:438.825) , CD14 (Fold Change:392.5985) , LGMN (Fold Change:354.5541) , S10 (Fold Change:291.7679)

Click the + icon to display details about each biomarker.

Click the - icon to remove the biomarker detail display.

When you click the + icon ( ) to pull down the list of biomarkers, you see two arrows next to each biomarker name. The arrows have the following meanings:

- The leftmost arrow indicates whether the gene in the signature or list is up-regulated (up arrow) or down-regulated (down arrow).
- The rightmost arrow (not shown above) indicates whether the gene in the comparison set is up-regulated (up arrow) or down-regulated (down arrow).



The leftmost arrow has meaning only for searches involving gene signatures or lists. The arrow is not shown for other searches.

Each analysis also includes the following download option:

- **Excel** – Download detailed analysis data (such as probe set, fold change ratio, p-value) to a Microsoft Excel spreadsheet.



## Study View

Click the **Study View** button to view the mRNA experiments that are returned and, optionally, *all* the analyses for each experiment – that is, those analyses that are considered statistically significant and those that are not.

Click the + icon to list all the analyses done for this experiment.

Click or hover the cursor over the experiment name or description to open details box.

Study result: 6 analyses from 4 experiment(s)

**GSE833: Amyotrophic lateral sclerosis**  
- 2 analyses found

DiseaseState => familial ALS vs control Excel

**BioMarkers** (top 5 of 1151):  
**CAPG** (Fold Change:858.1024) ↑, **AQP1** (Fold Change:438.825) ↑, **CD14** (Fold Change:392.5985) ↑, **LGMN** (Fold Change:354.5541) ↑, **S100A11** (Fold Change:291.7679) ↑

DiseaseState => sporadic ALS vs control Excel

**BioMarkers** (top 5 of 819):  
**CAPG** (Fold Change:221.8659) ↑, **FOXO4** (Fold Change:215.6599) ↑, **CD14** (Fold Change:193.2086) ↑, **PRKCSH** (Fold Change:190.8862) ↑, **PRKCD** (Fold Change:134.5789) ↑

**GSE4390: Analysis of expression in SOD1 transgenic mouse spinal cord**  
- 2 analyses found

### To drill down from the list of experiments:

1. Click the + icon (⊕) to the left of the experiment name to pull down a list of all the analyses done for the experiment.

The analysis list is similar to the list of the statistically significant analyses in the Analysis View. However, because Study View includes analyses ranked as not statistically significant, TEA scores and the designations co-regulated and anti-regulated are not specified for the analyses in Study View.

2. Click the + icon (⊕) to the left of the **BioMarker** label to pull down a list of applicable biomarkers for an analysis. Note that the same export options for biomarkers are available in Study View as in Analysis View.

### Export Results in Analysis View or Study View

While in either Analysis View or Study View, click the **Export Results** button to export the results data in the view to a Microsoft Excel spreadsheet:

Filters: Disease> Amyotrophic Lateral Sclerosis ⓧ [advanced](#) [save](#) [clear all](#)

mRNA Analysis (5, 6)

Show Filters Analysis View Study View **Export Results** ?

Study result: 6 analyses from 4 experiment(s)

The Export function writes the following information to the spreadsheet:

- Descriptions of each experiment returned from the search. This is the same information that appears in a details box for an experiment. In addition, associated diseases are exported to the Excel file.

- Information about the analyses associated with each experiment returned from the search. Information includes:
  - Analysis information displayed in the search results – for example, analysis description, TEA score, the list of matching biomarkers, and the probe set, fold change value, p-value, and TEA p-value associated with each biomarker.
  - Additional information about an analysis, such as QA criteria, analysis platform, descriptions of the biomarkers, biomarker type (such as gene expression), and associated diseases involved in the experiment.

All descriptions of experiments are written to one worksheet in the file, and all analysis data is written to a second worksheet in the file.

### Export Information about a Particular Analysis

To export details about all the biomarkers in a particular analysis, click the **Excel** button to the right of the analysis name – for example:

**About 4 results found**

Filters: **Disease> Amyotrophic Lateral Sclerosis** [advanced](#) [save](#) [clear all](#)

**mRNA Analysis (5, 6)**

Show Filters Analysis View Study View Export Results ?

**Study result: 6 analyses from 4 experiment(s)**

**GSE833: Amyotrophic lateral sclerosis**  
- 2 analyses found

DiseaseState => familial ALS vs control

**BioMarkers (top 5 of 1151):**  
CAPG (Fold Change:858.1024) ↑, AQP1 (Fold Change:438.825) ↑, CD14 (Fold Change:392.5985) ↑, LGMN (Fold Change:354.5541) ↑, S100A11 (Fold Change:291.7679) ↑

Excel

Note that the number of genes shown in parentheses after the **BioMarkers** label (1151 in the above example), which specifies the number of genes included in the analysis, may be less than the number of rows written to the spreadsheet. The Export function writes one row of data for each *probe set*, not each gene, and the same gene may be associated with multiple probe sets.

### Mouse Gene Homology in Search Results

Searches can return experiment results involving mouse genes. If experiment data is collected on a human gene and the corresponding mouse gene, a search against a human gene may potentially return results containing both human and mouse gene expression experiments.





For example, information on both can be found by clicking the **Export Results** button in the search results. The **Organism** column in the Excel worksheet indicates whether a particular measurement was made on a human gene or a mouse gene.

The following figure shows part of an Excel worksheet containing the results of a search against the MET gene:

| L  | M          | N                 | O            | P           | Q           | R      | S       | T           |
|--|------------|-------------------|--------------|-------------|-------------|--------|---------|-------------|
| Diseases   | Bio Marker | Description       | Organism     | ProbeSet    | Fold Change | RValue | p-Value | TEA p-Value |
|  | MET        | met proto-oncogen | HOMO SAPIENS | 203510_at   | 3.48        |        | 0.0008  | 0.0153      |
| Neoplasms by Histologic Type; Leukemia; Neoplasms; Leukemia, Myeloid   | Met        | met proto-oncogen | MUS MUSCULUS | 100309_at   | 3.72        |        | 0.0046  | 0.01828     |
|  | MET        | met proto-oncogen | HOMO SAPIENS | 1609_g_at   | -7.19       |        | 0.0036  | 0.01846     |
|  | Met        | met proto-oncogen | MUS MUSCULUS | 1422990_at  | 7.14        |        | 0.00004 | 0.02186     |
|  | Met        | met proto-oncogen | MUS MUSCULUS | 1422990_at  | -2.83       |        | 0       | 0.02559     |
| Acute Disease; Digestive System Diseases; Bacterial Infections and Mycoses; Infection; Sprains and Strains; Lung Diseases; Pathologic Processes; Bacterial Infections; | MET        | met proto-oncogen | HOMO SAPIENS | 213816_s_at | -2.7        |        | 0.00008 | 0.02849     |
|  | MET        | met proto-oncogen | HOMO SAPIENS | 203510_at   | -19.29      |        | 0.0049  | 0.02935     |
| Intestinal Neoplasms; Colorectal Neoplasms; Digestive System Diseases; Gastrointestinal Diseases; Neoplasms; Neoplasms by Site; Colonic Diseases; Intestinal Diseases  |            |                   |              |             |             |        |         |             |

### Additional Resources

An mRNA Analysis search result contains links to the following resources:

| Resource Link   | Description  |
|---|--|
| Experiment name<br>Example:<br> <a href="#">GSE5281</a>   | View information about the experiment, including title, description, and primary investigator.<br><br>The display may contain links to additional information, such as NCBI GEO and ArrayExpress data. |
| QA criteria<br>Example:<br> <a href="#">Compound:Time =&gt; 4 h -&gt; bacterial endotoxin vs control</a> | View key parameters of the experiment, such as p-Value and fold-change cutoffs, analysis platform, and methodology.  |
| Gene<br>Example:<br><a href="#">SOCS5</a>    | Search the following sites for information about the gene:<br><br><div> <a href="#">Entrez Gene</a> <a href="#">Entrez Global</a> <a href="#">Google Scholar</a> </div>                                |
|  <a href="#">Excel</a>   | Export data (such as gene, probe set, and fold-change ratio) for the matching biomarkers in a particular analysis to Microsoft Excel.  |

## Literature Tab

This result category contains hits from a set of curated articles within a set of selected diseases.

Click the **Literature** tab to display the results in this category. The following figure shows the controls that appear at the top of the results list.

These controls give you access to the following views and operations:

- **Show Filters** button – Define additional filters to further refine the search results.
- **Export Results** button – Write curation data to Microsoft Excel.
- **Results for** dropdown – Specify the type of literature results you want to see in the categories.

## Documents Tab

The search results in this category are based on internal text-indexed document repositories.

# TEA Analyses

---

Target Enrichment Analysis (TEA) measures the enrichment of a gene signature, gene list, or pathway in a microarray expression experiment.



For information on how TEA scores are calculated, see [Appendix A: How TEA Scores Are Calculated](#) on page 109.

## TEA Indicators Applied to Individual Biomarkers

The Study View of mRNA Analysis search result lists all experiments that satisfy the search criteria. Further, in Study View, you can list:

- All of an experiment's analyses that satisfy the search criteria
- All of an analysis' biomarkers that satisfy the search criteria

To drill down to the matching analyses in an experiment, click the **+** icon (⊕) next to the experiment name. To drill down to the matching biomarkers in an analysis, click the **+** icon next to the label **BioMarkers** under the analysis name.

The following example shows the experiment **GSE833** in Study View. The biomarkers for the analysis **DiseaseState => familial ALS vs control** and **DiseaseState => sporadic ALS vs control** are displayed:

**mRNA Analysis (5, 6)**

Show Filters Analysis View Study View Export Results ?

**Study result: 6 analyses from 4 experiment(s)**

**GSE833: Amyotrophic lateral sclerosis**  
- 2 analyses found

DiseaseState => familial ALS vs control Excel

**BioMarkers** (top 5 of 1151):  
CAPG (Fold Change:858.1024) ↑, AQP1 (Fold Change:438.825) ↑, CD14 (Fold Change:392.5985) ↑, LGMN (Fold Change:354.5541) ↑, S100A11 (Fold Change:291.7679) ↑

|         |   | ProbeSet    | Gene    | Fold Change | p-Value | TEA p-Value |
|---------|---|-------------|---------|-------------|---------|-------------|
| CAPG    | ↑ | M94345_at   | CAPG    | 858.102     | 0       | 0           |
| AQP1    | ↑ | M77829_s_at | AQP1    | 438.825     | 0.019   | 0.00295     |
| CD14    | ↑ | X13334_at   | CD14    | 392.598     | 0       | 0           |
| LGMN    | ↑ | D55696_at   | LGMN    | 354.554     | 0       | 0           |
| S100A11 | ↑ | D38583_at   | S100A11 | 291.768     | 0.037   | 0.00001     |

DiseaseState => sporadic ALS vs control Excel

**BioMarkers** (top 5 of 819):  
CAPG (Fold Change:221.8659) ↑, FOXO4 (Fold Change:215.6599) ↑, CD14 (Fold Change:193.2086) ↑, PRKCSH (Fold Change:190.8862) ↑, PRKCD (Fold Change:134.5789) ↑

Notice the rightmost column of biomarker values: **TEA p-Value**. These normalized p-values are intermediate values in the TEA calculation. To be considered a statistically significant analysis, an analysis must have at least one matching biomarker with a TEA p-Value of less than 0.05.

The following figure shows the same experiment and analysis from the figure above, but in Analysis View:

**mRNA Analysis (5, 6)**

Show Filters Analysis View Study View Export Results ?

**Analysis result: 5 analyses from 4 experiment(s)**

**GSE833 - DiseaseState => sporadic ALS vs control** Excel

**BioMarkers** (top 5 of 118):  
CAPG (Fold Change:221.8659) ↑, FOXO4 (Fold Change:215.6599) ↑, CD14 (Fold Change:193.2086) ↑, PRKCSH (Fold Change:190.8862) ↑, PRKCD (Fold Change:134.5789) ↑

|        |   | ProbeSet       | Gene   | Fold Change | p-Value | TEA p-Value |
|--------|---|----------------|--------|-------------|---------|-------------|
| CAPG   | ↑ | M94345_at      | CAPG   | 221.866     | 0       | 0           |
| FOXO4  | ↑ | X93996_rna1_at | FOXO4  | 215.66      | 0       | 0           |
| CD14   | ↑ | X13334_at      | CD14   | 193.209     | 0       | 0           |
| PRKCSH | ↑ | U50327_s_at    | PRKCSH | 190.886     | 0       | 0           |
| PRKCD  | ↑ | D10495_at      | PRKCD  | 134.579     | 0       | 0           |

**GSE833 - DiseaseState => familial ALS vs control** Excel

**BioMarkers** (top 5 of 104):  
CAPG (Fold Change:858.1024) ↑, AQP1 (Fold Change:438.825) ↑, CD14 (Fold Change:392.5985) ↑, LGMN (Fold Change:354.5541) ↑, S100A11 (Fold Change:291.7679) ↑

Statistically significant analyses are candidates for display in the Analysis View, after further TEA calculations are performed to determine whether the analysis is a **significant TEA analysis** or an **insignificant TEA analysis**.

## TEA Indicators Applied to an Analysis

The TEA algorithm assigns an aggregate score to each analysis within an experiment. A TEA score is a binomial distribution of normalized p-values, calculated in the context of the following factors:

- **With gene signatures and gene lists** – The level of co-regulation or anti-regulation of the genes within the gene signature or gene list, as compared with the experiment.
- **With pathways** – The level of up-regulation or down-regulation of the genes within the pathway, as compared with the experiment.



For details on the TEA algorithm, see [Appendix A: How TEA Scores Are Calculated](#) on page 109.

TEA identifies experiments where the genes in the signature, list, or pathway are *differentially modulated*, indicating that the target is affected by the treatment, disease or other topic examined in the experiment.

## What the TEA Score Means

The TEA score displayed for an analysis of an experiment is not the actual TEA score calculated by the TEA algorithm. TEA scores are typically very small decimal numbers that are not easily human-readable. To aid users in interpreting the relative value of TEA scores, scores are converted to a larger number, as follows:

$$\text{Displayed\_TEA\_Score} = -\log(\text{Actual\_TEA\_Score})$$

The larger the displayed TEA score, the more statistically significant is the analysis.

Typically, displayed TEA scores for statistically significant analyses of experiments range from 3 to 30 or 40.

Analyses of experiments are grouped into the categories **Significant TEA Analyses** and **Insignificant TEA Analyses**, as follows:

- Significant TEA analyses have a displayed TEA score of  $\geq 2.9957$ .
- Insignificant TEA analyses have a displayed TEA score of  $< 2.9957$ .

## What Co-/Anti-Regulation and Up-/Down-Regulation Mean

An analysis of a statistically significant experiment returned from a search against a gene signature or list is designated as *co-regulated* or *anti-regulated*. An analysis of a statistically significant experiment returned from a search against a pathway is designated as *up-regulated* or *down-regulated*.

The following table describes what these terms imply in the context of an analysis of a statistically significant experiment:

|                       | Gene Signature/List  | Pathway   |
|-----------------------|--|---|
| <b>Co-Regulated</b>   | Genes that are up-regulated in the signature or list are predominantly up-regulated in the experiment.<br><br>Genes that are down-regulated in the signature or list are predominantly down-regulated in the experiment. | n/a   |
| <b>Anti-Regulated</b> | Genes that are up-regulated in the signature or list are predominantly down-regulated in the experiment.<br><br>Genes that are down-regulated in the signature or list are predominantly up-regulated in the experiment. | n/a   |
| <b>Up-Regulated</b>   | n/a  | Genes in the experiment are predominantly up-regulated.   |
| <b>Down-Regulated</b> | n/a  | Genes in the experiment are predominantly down-regulated. |

## TEA Indicators Applied to an Individual Gene

In an analysis list, TEA indicators for a gene appear as arrows, as shown in the figure below. The leftmost arrow represents the gene expression in the gene signature or list. The rightmost arrow represents the gene expression in the experiment:

**BioMarkers** (3 signature/pathway genes matched):

|       |   |                       |                    |                      |                      |
|-------|---|-----------------------|--------------------|----------------------|----------------------|
| GBP1  |   | ProbeSet: 202270_at   | Gene: <b>GBP1</b>  | Fold Change: -1.9417 | TEA p-Value: 0.00015 |
| CASP1 |   | ProbeSet: 211367_s_at | Gene: <b>CASP1</b> | Fold Change: -1.7762 | TEA p-Value: 0.00029 |
| CASP1 |   | ProbeSet: 206011_at   | Gene: <b>CASP1</b> | Fold Change: -1.5244 | TEA p-Value: 0.00075 |



The leftmost arrow appears only for gene signatures and gene lists.

The direction of the arrows indicates the following:

- **Up-arrow** – An upward-pointing arrow alongside a gene indicates that the gene is up-regulated in the gene signature/list (leftmost arrow) or in the experiment (rightmost arrow).

If both arrows point in the same direction, the gene is co-regulated in the signature/list and the experiment. If the arrows point in opposite directions, the gene is anti-regulated.

- **Down-arrow** – A downward-pointing arrow alongside a gene indicates that the gene is down-regulated in the gene signature/list (leftmost arrow) or in the experiment (rightmost arrow).

If both arrows point in the same direction, the gene is co-regulated in the signature/list and the experiment. If the arrows point in opposite directions, the gene is anti-regulated.

The relationships between TEA indicators for genes and TEA indicators for an experiment are as follows:

- **Co-regulated genes** – Up- or down-regulated genes in the signature/list are similarly up- or down-regulated in the experiment.
- **Anti-regulated genes** – Up- or down-regulated genes in the signature/list are conversely down- or up-regulated in the experiment.



## Chapter 3

# Dataset Explorer

Dataset Explorer lets you compare data generated for test subjects in two different study groups, based on criteria and points of comparison that you specify. Dataset Explorer is useful to help you test a hypothesis that involves the criteria and points of comparison that you select.

## Overview of the UI

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The figure below shows the Dataset Explorer interface. It is divided into two panes:

### Left pane

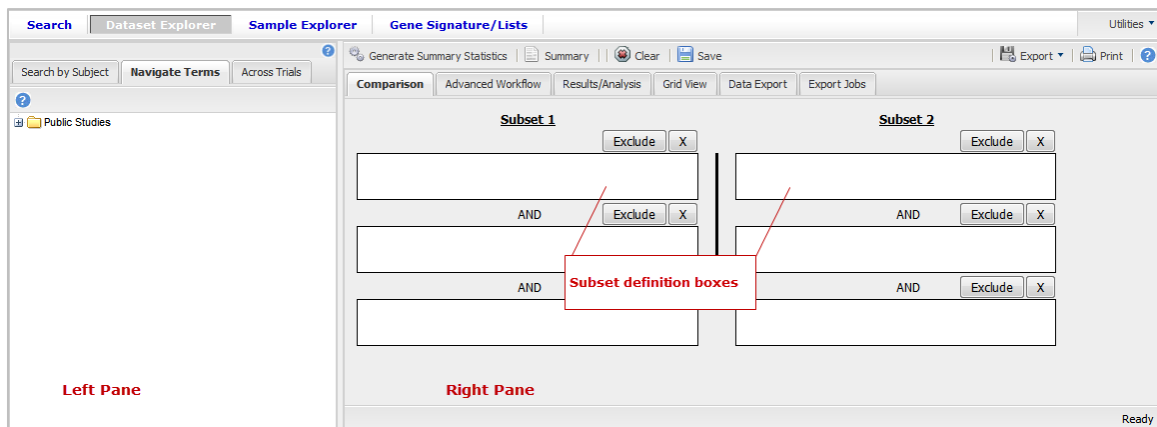
- Lets you select the study of interest.
- Provides a Microsoft Windows Explorer-like navigation tree where you select the criteria for membership in the study groups and the points of comparison between the study groups.

### Right pane

- Lets you define the criteria that test subjects must satisfy to become members of one of the two groups being compared. Each of these groups is called a *subset* because it typically contains only some of the subjects in the actual study group involved in the study.

You define the criteria for the subsets in the subset definition boxes shown below. Subjects who do not satisfy the criteria you define are excluded from the subsets.

- Provides summary data about the subjects being compared, and several different views of the comparison data.



The following table describes the buttons and tabs in the right pane of Dataset Explorer:

| Button or Tab                      | Description  |
|------------------------------------|--|
| Generate Summary Statistics button | Displays tables and charts that describe demographic information about the subjects in the subsets, and also analyses of criteria included in the subset definitions.<br><br>The tables and charts are displayed in the Results/Analysis view.                               |
| Summary button                     | Displays a summary of the query criteria you specified. Dataset Explorer uses these criteria to select the subjects for the subsets.   |
| Clear button                       | Clears all the criteria in the subset definition boxes.  |
| Save button                        | Saves the criteria definition. This allows you to re-generate the comparison at a later time without having to reconstruct the criteria that select the subjects for the subsets.<br><br>For more information, see <a href="#">Saving Comparison Definitions</a> on page 39. |
| Export button                      | Exports summary statistics data or expression data to Microsoft Excel.   |
| Print button                       | Prints the tables and charts in the Results/Analysis view.   |
| Comparison tab                     | Removes the currently displayed view (that is, the Results/Analysis view, or Grid view) and re-displays the subset definition boxes. This allows you to further refine the subjects for the comparison.  |
| Advanced Workflow tab              | Displays advanced analyses and visualizations that you can perform on data.  |
| Results/Analysis tab               | Displays tables and charts containing comparison and analysis data generated from the "Summary Statistics" workflow.   |
| Grid View tab                      | Displays the comparison and analysis data in grid format.  |
| Jobs tab                           | Displays previously run analyses.  |
| Data Export tab                    | Allows you to select data to export for further analysis in an external tool.  |
| Export Jobs tab                    | Displays previously exported jobs.   |

## Using Dataset Explorer

Four basic tasks are involved in using Dataset Explorer:

- Select the study (clinical trial or experiment) to use in the comparison.
- Specify the criteria for membership in the two study groups. Note some analyses only allow for the specification of one group at this time.
- Generate summary statistics for the two study groups.
- Specify the points of comparison to apply to the study groups.



You may see the notations **NA** and **Unknown** in the study data. **NA** indicates not applicable, and **Unknown** indicates not available.

### Public and Private Studies

Dataset Explorer studies can be either public or private. Public studies can be found both in the **Public Studies** folder of the Dataset Explorer navigation tree, as well as in the research-specific folders.

You can perform all the operations described in this chapter on public studies. No special privileges are required.

To perform operations described in this chapter on a private study, a tranSMART Administrator must assign you access rights to the study. Access rights are based on the following access levels:

| Access Level | Privileges  |
|--------------|---|
| VIEW         | Define the criteria for the study groups to be compared, generate summary statistics for the study groups, and specify points of comparison for the study groups. |
| EXPORT       | All privileges of the VIEW access level, plus the ability to export comparison data or expression data to a Microsoft Excel spreadsheet.                          |
| OWN          | All VIEW and EXPORT privileges.<br><br>This access level can only be assigned to the owner of the study.  |

If you do not have access rights to the study you want (that is, if the study name is grayed out), contact a tranSMART Administrator. The administrator will contact the study owner to find out if you should be granted VIEW access, EXPORT access, or no access.



Even if you have no access rights to a private study, you can read a description of the study. For information, see [Viewing a Study](#) on page 45.

## Selecting the Study

You select the study in the left pane of Dataset Explorer. You have several ways to select the study, based on the tab you choose – Search by Subject or Navigate Terms.

### Search by Subject Tab

Use this tab to search for studies using one or a combination of the following fields:

- **Search** field. Lets you specify part or all of a term from a study that is stored in the tranSMART database. Search terms may include part or all of a study name, the text in a branch of the Dataset Explorer navigation tree, or some attribute of a study, such as a disease or an area of clinical interest.

Example:

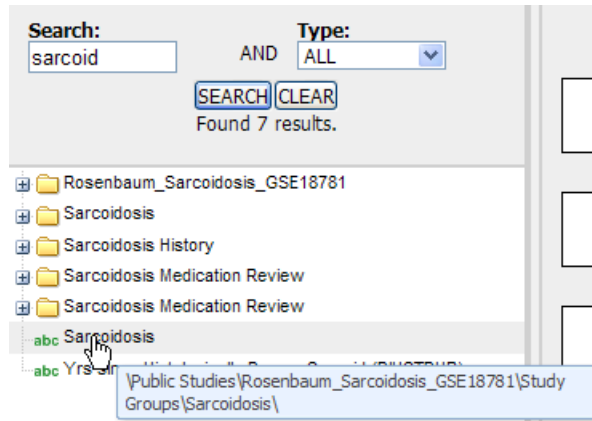
The screenshot shows the 'Search by Subject' tab selected. The search bar contains the text 'acute'. The 'Type' dropdown is set to 'ALL'. The 'SEARCH' button is highlighted. Below the search bar, it says 'Found 1 results.' and lists a single result: 'Acute\_Physiologic\_Hyperinsulinemia\_Coletta\_GSE9105'.

If you want to base your search on a study name, note the following naming conventions for Public Studies in Dataset Explorer:

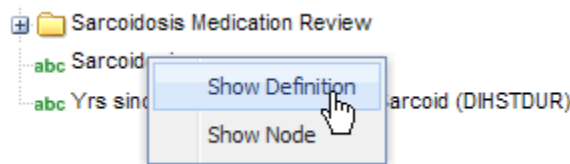
| Study Type     | Naming Convention   |
|----------------|---|
| Public Studies | <p>Name segments in the following typical format:<br/> <i>Condition_StudyFirstAuthor_GEOid</i></p> <p>Example: ProstateCancer_Ambs_GSE6956</p> <p>If you prefer, you can rearrange the order of the segments (for example, author segment first). The name structure is determined by the ETL process that loads the data into the i2b2 database.</p> |

## Selecting and Opening a Study in a Search Result

A search result may include multiple entries. Further, an entry may not indicate the study it is from. To see the name of the study that an entry represents, hover the mouse pointer over the entry – for example:



If you want more information about the study represented by an entry, right-click the entry, then click **Show Definition** to open the details box for the study:



To open a study from an entry in a search result, right-click the entry, then click **Show Node**. The study appears in the Dataset Explorer navigation tree, where you can open any of the branches (nodes) in the study.



You may need to scroll down slightly in the navigation tree to see the study.

## Navigate Terms Tab

Use this tab to browse through all the experiments in the navigation tree to select and open the study you want.

Studies that are grayed out are private studies that you are not authorized to access.

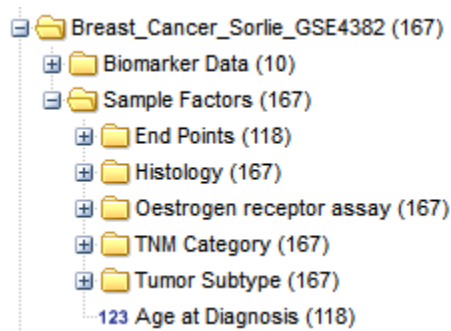
To display the details box for a study, right-click the study name and click **Show Definition**. You can display the details box for a study whether or not the study is grayed out.

## Branches and Leaves of the Navigation Tree

The Dataset Explorer navigation tree looks and works much like the Microsoft Windows Explorer. Windows Explorer is a hierarchy of folders, sub-folders, and files. Dataset Explorer is a hierarchy of folders and sub-folders (the branches) and values (the leaves) that reflect aspects of the trial, such as research metrics, compounds used, and patient demographics.

In Dataset Explorer, all levels of the tree, both branches and leaves, are referred to as nodes.

The following figure shows typical top-level nodes of a study. Some studies may not require all of these nodes, and others may require additional nodes (such as Published Conclusions):



The following table describes possible top-level nodes of a study:

| Node                   | Description   |
|------------------------|---|
| Biomarker Data         | Measurements of biomarkers such as RBM antigens, gene expressions, antibodies and antigens in ELISA tests, and SNPs.  |
| Clinical Data          | Primary and secondary endpoints, and other measurements from the study.   |
| Samples and Timepoints | Tested samples (such as tissue or blood) and time periods when the samples were taken.  |
| Scheduled Visits       | Periodic stages of the trial during which patients are seen.  |
| Design Factors         | Compounds involved in the study, dosages, and regularity with which the compounds were administered.<br><b>Note:</b> With clinical trials, this node is typically named Treatment Groups. |
| Sample Factors         | Patient information, such as demographics and medical history.  |

## Populating the Study Groups

You populate the study groups by defining criteria that members of each group must satisfy. For example, members of study groups might be required to satisfy a weight or age requirement. Dataset Explorer lets you build a set of criteria for each study group that can be as simple or as complex as you need.

The study groups you define are called *subsets*, because typically, after your criteria are applied, the members of a resulting study group are a subset of the full study group that participated in the study.

## Selecting Criteria for the Study Groups

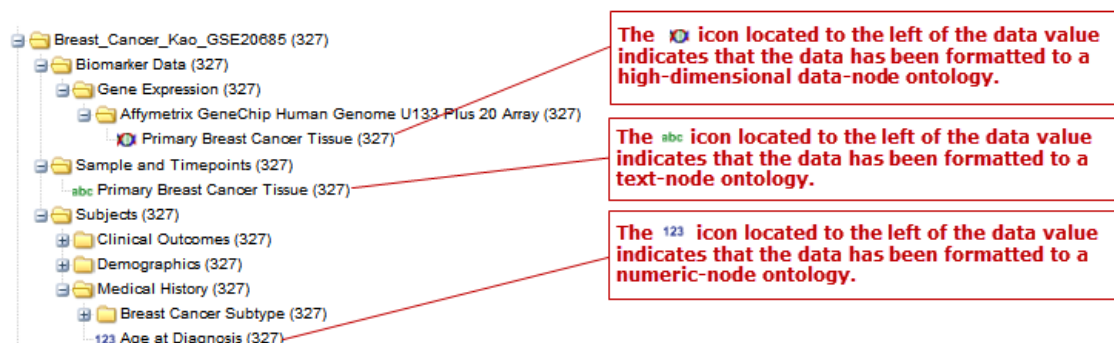
You define the study groups by selecting criteria (called concepts) from the navigation tree and dragging them into the subset definition boxes.

### Visual Aids to Help You Select the Criteria

Each concept node in the navigation tree displays the following information about the concept:

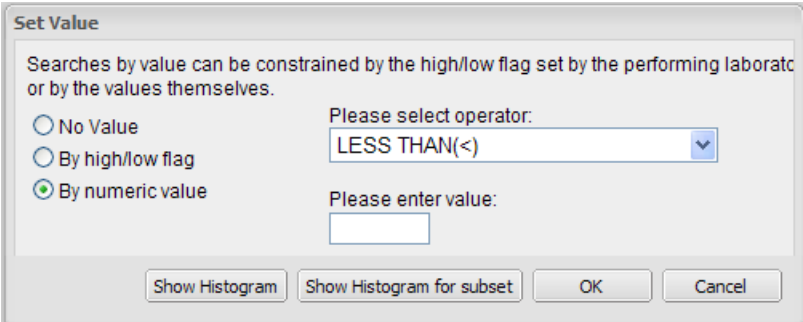
- The numbers in parentheses at each node of the tree indicate the number of subjects to whom that node applies. For example, in the figure below, there are a total of 327 subjects in the study.
- In tranSMART, data values are represented in one of three ways: by number, by text, or by high dimensional data (SNP, gene expression, etc.) stored as *arrays*.

The three types of data values are described in the illustration below:



### Specifying a Numeric Value

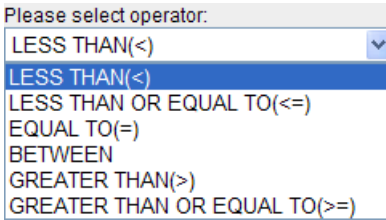
When you drag a numeric concept into a subset definition box, the Set Value dialog appears:

The image shows a 'Set Value' dialog box. At the top, it says 'Set Value'. Below that, a paragraph reads: 'Searches by value can be constrained by the high/low flag set by the performing laboratory or by the values themselves.' There are three radio buttons: 'No Value', 'By high/low flag', and 'By numeric value'. The 'By numeric value' option is selected. To the right of the radio buttons, there is a label 'Please select operator:' followed by a dropdown menu showing 'LESS THAN(<)'. Below that, there is a label 'Please enter value:' followed by an empty text input field. At the bottom of the dialog, there are four buttons: 'Show Histogram', 'Show Histogram for subset', 'OK', and 'Cancel'.

Use the Set Value dialog to specify how you want to constrain the numeric values to use in the subset definition. To do so, first select one of the following choices:

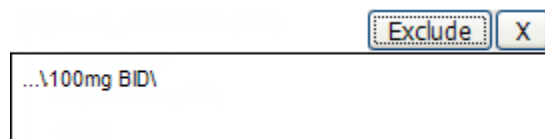
| Selection        | Description   |
|------------------|---|
| No Value         | Values are not constrained. All the numeric data associated with the concept are factored into the subset definition.<br><br>If you select <b>No Value</b> , no other information is required. Click <b>OK</b> to add the concept with all its associated numeric data to the subset.       |
| By high/low flag | If the testing laboratory has grouped the numeric values into high/low/normal ranges, select the range to factor into the subset definition.<br><br>When you select <b>By high/low flag</b> , the <b>Please select range</b> field appears. Select the range you want and click <b>OK</b> . |



| Selection        | Description  |
|------------------|--|
| By numeric value | <p>Values are constrained by an exact value or a range of values.</p> <p>After you select <b>By numeric value</b>:</p> <ul style="list-style-type: none"> <li>■ Select one of the following numeric operators in the <b>Please select operator dropdown</b>:<br/>  </li> <li>■ In <b>Please enter value</b>, type the numeric value that the operator applies to.<br/> For example, to constrain the ages of subjects to 50 years or younger, select LESS THAN OR EQUAL TO (&lt;=) in the dropdown, then type 50 in the <b>Please enter value</b> field.</li> <li>■ Click <b>OK</b>.</li> </ul> <p>See the next section for information on viewing the numeric values associated with the concept and that you can select from.</p> |



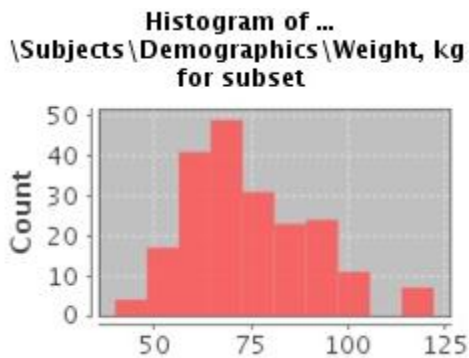
When finished defining the numeric constraint on the Set Value dialog, be sure to click **OK** and not press the **Enter** key. Pressing **Enter** will activate the subset button that has focus – the **Exclude** button in the example below:



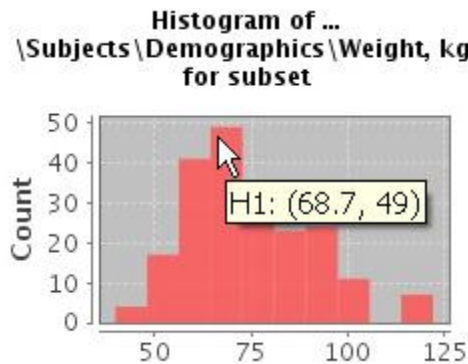
### Viewing the Numeric Values Associated with a Concept

Note the buttons **Show Histogram** and **Show Histogram for subset** in the Set Value dialog. The histograms show how the numeric values associated with the concept that you placed in the subset box are distributed among the subjects across both subsets, or in the particular subset you are currently defining, respectively.

A histogram may be helpful in determining the number to set as the constraining factor for a concept. For example, suppose you drag a Weight concept into a subset box, then click **Show Histogram for subset**. In the following histogram of the weights of test subjects, the weights range from about 25 kg to just under 125 kg. The largest bin represents just under 50 subjects. You may want to use these weight parameters to help you determine the value to set for the weight concept.



You can get more specific information about the number of subjects represented by a particular bin and the average of the values in the bin by hovering the mouse cursor over the bin you are interested in. For example, in the following figure, the largest bin represents 49 subjects with an average weight of 68.7 kg:

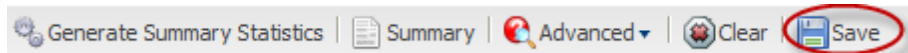


## Saving Comparison Definitions

You may save your search criteria in order to regenerate the comparison at a later time without having to redefine the subsets.

### To save search criteria:

1. Run tranSMART, then click the **Dataset Explorer** tab.
2. Select the study of interest.
3. Define the cohorts whose data points will be represented.
4. Click **Save**:



5. Click **Email this comparison**:



Your email application will open with a link to the saved comparison.

6. Send the email to yourself so that you can retrieve the comparison later. Optionally, send it to colleagues who might be interested in the comparison.

When you or someone else clicks the link in the email, Dataset Explorer opens with the subset boxes pre-defined.

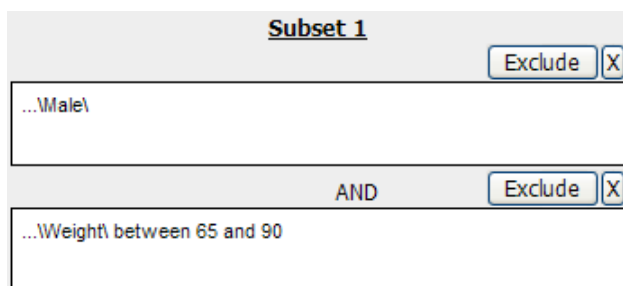
## Joining Multiple Criteria for a Subset Definition

Multiple criteria for a subset definition are joined by one of the following logical operators: AND, OR, or AND NOT.

The rules for joining multiple criteria are as follows:

- Criteria in separate subset definition boxes are joined by an AND operator.

For example, the following definition boxes select only male subjects, AND males whose weights are between 65 kg and 90 kg:



- Criteria within the same subset definition box are joined by an **OR** operator.

For example, to use the extreme ends of the weight scale for your weight criterion, you might add the following to a definition box:

The image shows a software interface for defining a subset. At the top, it is labeled "Subset 1". To the right of the label are two buttons: "Exclude" and "X". Below the label is a text box containing two lines of criteria: "...Weight\ <=50" and "...Weight\ >=100".

This criterion selects subjects whose weight is either 50 kg or less, **OR** 100 kg or greater.

- To join a definition box with an **AND NOT** operator, click the **Exclude** button above the definition box.


The figure below selects only male subjects, but not those who weigh between 50 kg and 100 kg:

The image shows a software interface for defining a subset. At the top, it is labeled "Subset 1". To the right of the label are two buttons: "Exclude" and "X". Below the label is a text box containing the criterion "...Male\". Below this text box is a horizontal line with the word "AND" in the center. To the right of "AND" are two buttons: "Include" and "X". Below this line is another text box containing the criterion "...Weight\ between 50 and 100".

Note that when you click the **Exclude** button, the button label changes to **Include**, allowing you to join the criteria in the box with an **AND** operator later if you choose.

## Modifying or Deleting Criteria

To delete or modify a criterion in a subset definition box, right-click the criterion and select either **Delete** or **Set Value**.

To remove the entire contents of a subset definition box from the subset definition, click the **X** icon () above the box:

The image shows a software interface for defining a subset. At the top, it is labeled "Subset 1". To the right of the label are two buttons: "Exclude" and "X". The "X" button is circled in red. Below the label is an empty text box.

## Generating Summary Statistics

When you finish defining criteria for the groups to compare – the subsets – click the **Generate Summary Statistics** button.

tranSMART displays tables and charts of information that describe the subsets. The information is displayed in the Results/Analysis view in the following sections:

- A summary of the criteria used to define subsets to compare. Example:

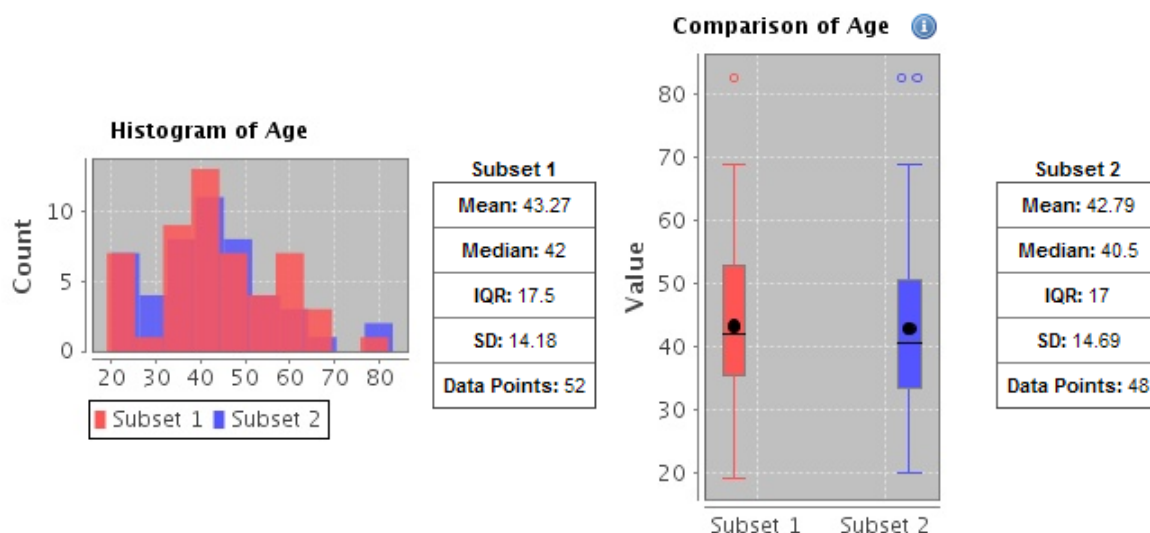
| Query Summary for Subset 1  | Query Summary for Subset 2  |
|---|---|
| (\\Public Studies\\Public Studies\\Lymphoma_Staudt_GSE10846\\Biomarker Data\\Gene Expression\\ )<br>AND<br>(\\Public Studies\\Public Studies\\Lymphoma_Staudt_GSE10846\\Subjects \\Demographics\\Gender\\Female\\ ) | (\\Public Studies\\Public Studies\\Lymphoma_Staudt_GSE10846\\Biomarker Data\\Gene Expression\\ )<br>AND<br>(\\Public Studies\\Public Studies\\Lymphoma_Staudt_GSE10846\\Subjects \\Demographics\\Gender\\Male\\ ) |

- A table showing the number of subjects in each subset who match the subset criteria. Example:

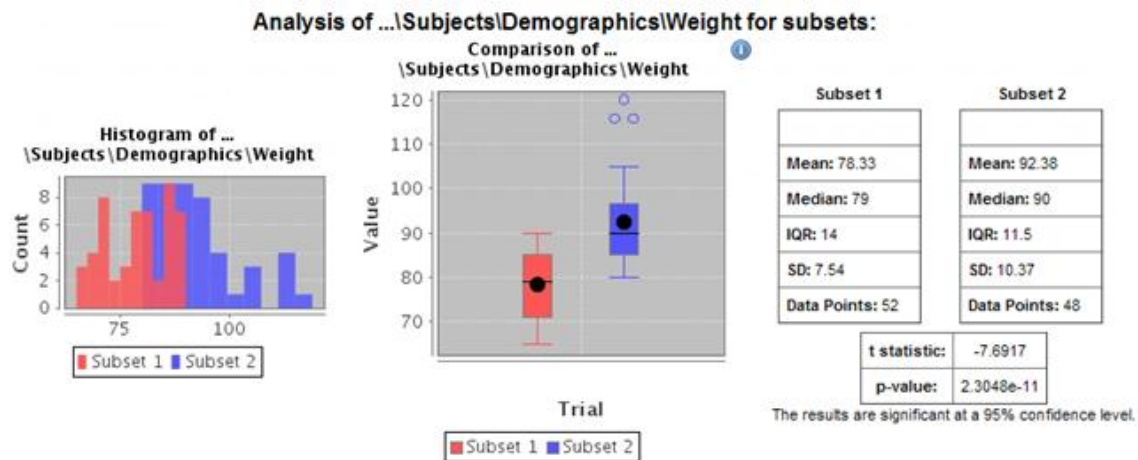
| Subject Totals |      |          |
|----------------|------|----------|
| Subset 1       | Both | Subset 2 |
| 52             | 25   | 48       |

In this example, 52 subjects matched the criteria for Subset 1, and 48 matched the criteria for Subset 2. Further, 25 subjects matched the criteria for both subsets (and thus, were included in both).

- Tables and charts that show how the subjects who match the criteria fit into age, sex, and race demographics. Example (showing the age portion only):



- Analyses of the concepts you added to the subsets from the navigation tree.  
Example (showing the weight concept):



## Significance Tests

The above figure includes the results of significance testing that Dataset Explorer performs:

|              |            |
|--------------|------------|
| t statistic: | -7.6917    |
| p-value:     | 2.3048e-11 |

The results are significant at a 95% confidence level.

Significance testing is designed to indicate whether the reliability of the statistics is 95% or greater, based on p-value.

Dataset Explorer calculates the significance result using either t-test or chi-squared statistics to determine the p-value:

- For continuous variables (for example, subject weight or age), a t-test compares the observed values in the two subsets.

tranSMART uses the following Java method to calculate the t-test statistic:

[http://commons.apache.org/math/apidocs/org/apache/commons/math/stat/inference/TTest.html#tTest\(double\[\],%20double\[\]\)](http://commons.apache.org/math/apidocs/org/apache/commons/math/stat/inference/TTest.html#tTest(double[],%20double[]))

- For categorical values (for example, diagnoses), a chi-squared test compares the counts in the two subsets.

tranSMART uses the following Java method to calculate the chi-squared statistic:

[http://commons.apache.org/math/apidocs/org/apache/commons/math/stat/inference/ChiSquareTest.html#chiSquareTest\(long\[\]\[\]\)](http://commons.apache.org/math/apidocs/org/apache/commons/math/stat/inference/ChiSquareTest.html#chiSquareTest(long[][]))

If there is not enough data to calculate a test, Dataset Explorer displays a message indicating the insufficient quantity data. Also, significance test results are not displayed in the following circumstances:

- If two identical subsets are defined. In this case, the significance test results are not meaningful.
- If all subjects in the first subset have one set of values for the categorical value, and all subjects in the second subset have other categorical values. For example, suppose you set Subset 1 to contain only males and Subset 2 to contain only females. Also, suppose that Subset 1 has 15 subjects and Subset 2 has 20. If you then try to show statistics by gender, a table like the following would result:

|        | Subset 1 | Subset 2 |
|--------|----------|----------|
| Female | 0        | 20       |
| Male   | 15       | 0        |

In this case, the chi-squared function doesn't return meaningful results.

## Defining Points of Comparison

Once you establish the subsets of subjects that you want to compare, you can apply one or more points of comparison to the subsets.

A point of comparison is a concept in the navigation tree.

### To apply a point of comparison to the subsets:

1. You must already have defined the subsets and have generated summary statistics for the subsets, as described in the previous section.
2. Drag the concept that you want to introduce as the point of comparison from the navigation tree, and drop it anywhere in the Results/Analysis view.

As soon as you drop the point of comparison into the Results/Analysis view, tranSMART begins to compare the subsets based on that point of comparison. When finished, tranSMART displays a side-by-side summary of how the subjects in each subset match or respond to the point of comparison.

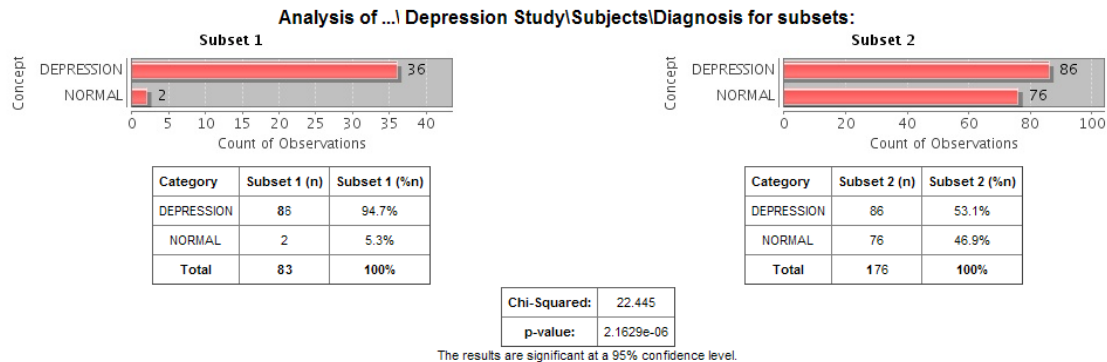
## Results of a Comparison

In a comparison of subjects in a psychological study, suppose Subset 1 contains subjects with a substance abuse problem, and Subset 2 contains subjects with no substance abuse assessment.

After the subsets are defined and summary statistics are generated, a diagnosis of depression is dropped into the Results/Analysis view as a point of comparison.

tranSMART displays a side-by-side comparison of the subjects in each subset, indicating that almost all the subjects with a substance abuse problem have been diagnosed with depression, while that diagnosis for those with no substance abuse problem is more evenly split.

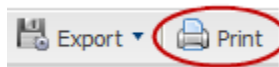
The comparison is placed at the top of the Results/Analysis view, above the demographic definitions plus any other earlier comparisons:



To keep the size of the preceding figure within production limits, the demographics (age, sex, and race) portions of the figure have been excluded.

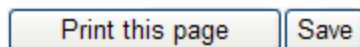
## Printing or Saving the Contents of the Results/Analysis View

1. With the Results/Analysis view displayed, click **Print**:



The entire contents of the Results/Analysis view appear in a separate browser window.

2. Click one of the following buttons at the top of the browser window:



## Copying Individual Charts in the Results/Analysis View

If you are interested in a particular chart in the Results/Analysis View, you can copy the chart to a file, as follows:

1. With the Results/Analysis view displayed, click **Print**.

The entire contents of the Results/Analysis view appear in a separate browser window.



2. Right-click the chart to copy.
3. In the Internet Explorer popup menu, click **Save Picture As**.
4. In the Save Picture dialog, specify the name, location, and the file type for the chart.
5. Click **Save**.

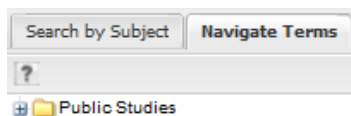
## Viewing a Study

---

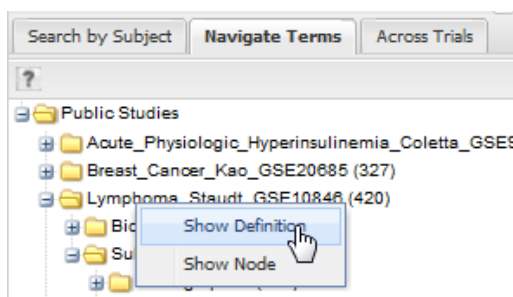
You can view a description of any Dataset Explorer study, whether or not you have access rights to the study.

### To view a description of a study:

1. In Dataset Explorer, click the **Navigate Terms** tab.
2. Open the top-level node for the list of studies you are interested in – for example, click the **+** icon (⊕) next to Public Studies to open the list of experiments:



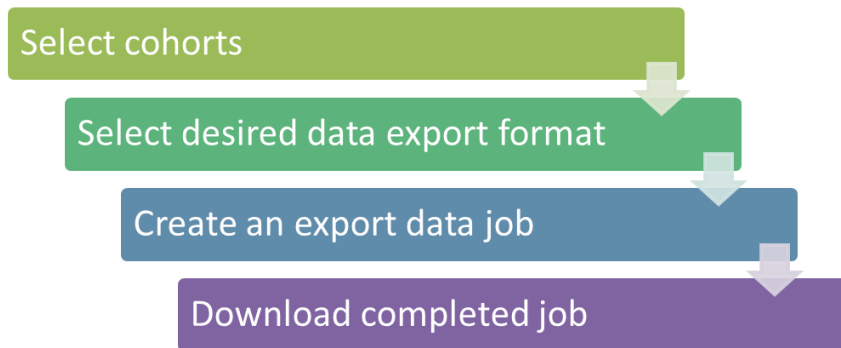
3. Right-click the particular study you are interested in.
4. Click the **Show Definition** popup:



The Show Concept Definition dialog appears, showing the title, description, and other information about the study.

## Exporting Dataset Explorer Findings

The Data Export tab allows you to export your data locally for further analysis in several different formats. Exporting data using this tool involves the following high-level tasks:



Supported file formats include:

- Clinical and low dimensional biomarker data
- Gene expression data
- SNP data
- Gene set enrichment analysis (GSEA)



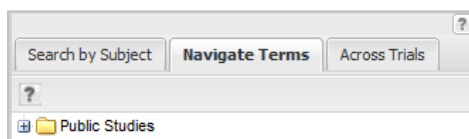
For more information on GSEA data files, visit the following site:

[http://www.broadinstitute.org/cancer/software/gsea/wiki/index.php/Data\\_formats](http://www.broadinstitute.org/cancer/software/gsea/wiki/index.php/Data_formats)

### To export Dataset Explorer findings to your local machine:

1. Click the tranSMART **Dataset Explorer** tab to display the Dataset Explorer window.
2. In the left pane of the Dataset Explorer window, click the **Navigate Terms** tab.

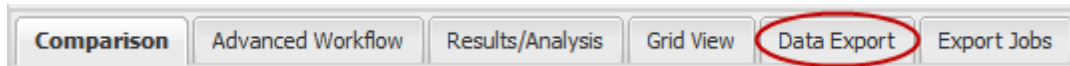
The navigation tree appears, showing the categories of available studies:



3. Select the study of interest.
4. Define the cohorts whose data points are of interest.

Now that the subsets are defined, you are ready to export data from the study that applies to the subsets.

5. Click the **Data Export** tab:



The Data Export page appears with your selected cohorts:

Comparison Advanced Workflow Results/Analysis Grid View **Data Export** Export Jobs

**Instructions:** Select the check boxes to indicate the data types and file formats that are desired for export. Then, click on the "Export Data" button at the bottom of the screen to initiate an asynchronous data download job. To download your data navigate to the "Export Jobs" tab.

|   | Subset 1  | Subset 2  |
|---|---|---|
| Selected Cohort                           | (Public Studies\Breast_Cancer_Sorlie_GSE4382\Sample F...<br>AND<br>(Public Studies\Breast_Cancer_Sorlie_GSE4382\Sample F...   | (Public Studies\Breast_Cancer_Sorlie_GSE4382\Sample F...<br>AND<br>(Public Studies\Breast_Cancer_Sorlie_GSE4382\Sample F...   |
| Clinical & Low Dimensional Biomarker Data | Data is available for 8 patients<br>Export (.TXT) <input type="checkbox"/><br><br>Metadata will be downloaded in a separate file.   | Data is available for 36 patients<br>Export (.TXT) <input type="checkbox"/><br><br>Metadata will be downloaded in a separate file.  |
| Gene Expression Data                      | Raw Data is not available.  | Processed Data is available for<br>Stanford cDNA microarrays: 1 patients<br>Export (.TXT) <input type="checkbox"/><br><br>Raw Data is not available.<br><br>Metadata will be downloaded in a separate file. |
| Gene Set Enrichment Analysis (GSEA)       | Processed Data (for both subsets) is available for<br>Stanford cDNA microarrays: 1 patients<br>Export (.GCT & .CLS) <input type="checkbox"/><br><br>Metadata will be downloaded in a separate file. |   |

Export Data

6. Select the check boxes to indicate the data types and file formats that are desired for export.
7. Click **Export Data** at the bottom of the tranSMART browser window.

The command will now start a job. You may choose to have the job run in the background in order to continue with other analyses and cohort selection while the job completes. The job could take several minutes depending on the amount of data selected.

8. Click the **Export Jobs** tab to access completed jobs or to check the status of a pending job.

Jobs follow the naming convention *User - Type of Job Run - Job ID*.

9. Click the name of the job you processed:

Comparison

Advanced Workflow

Results/Analysis

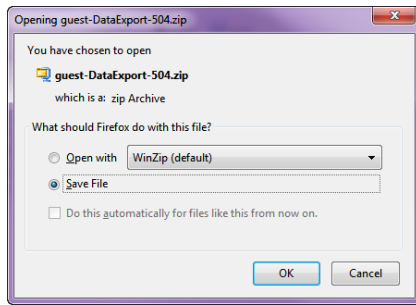
Grid View

Data Export

Export Jobs

| Name                 | Query Summary | Status         |
|----------------------|---------------|----------------|
| guest-DataExport-504 |               | Gathering Data |
| guest-DataExport-503 |               | Completed      |
| guest-DataExport-502 |               | Completed      |

The Open File dialog box appears:



10. Select **Save File**, then click **OK**.

Your file will be sent to the **Downloads** folder on your local machine in a .zip file. The .zip file contains separate folders for subsets, clinical data, gene expression data, and other factors you may have specified during cohort selection.

## Generating Advanced Analyses and Visualizations

Advanced analyses and visualizations offered with tranSMART allow a user to produce the following within Dataset Explorer:

- Heatmaps
  - [Standard Heatmap](#) (page 49)
  - [Hierarchical Clustering](#) (page 51)
  - [K-Means Clustering](#) (page 54)
  - [Marker Selection](#) (page 57)
- Advanced Analyses
  - [Box Plot with ANOVA](#) (page 60)
  - [Correlation Analysis](#) (page 62)
  - [Line Graph Analysis](#) (page 64)
  - [Principal Component Analysis](#) (page 65)
  - [Scatter Plot with Linear Regression](#) (page 68)
  - [Survival Analysis](#) (page 70)
  - [Table with Fisher Test Analysis](#) (page 73)

Dataset Explorer uses the R software environment for statistical computing and to generate analyses and visualizations. For more information, visit <http://www.r-project.org>.

## Generating Heatmaps

In Dataset Explorer, a heatmap is a matrix of data points for a particular set of biomarkers, such as genes, at a particular point in time and/or for a particular tissue sample in the study, as measured for each subject in the study.

In a Dataset Explorer heatmap, the biomarkers appear in the y axis, and the subjects appear in the x axis.



A heatmap can display data points for up to 1000 samples.

Dataset Explorer uses the R software environment for statistical computing and to generate analyses and visualizations. For more information, visit <http://www.r-project.org>.

You can generate the following types of heatmaps:

- [Standard Heatmap](#) (below)
- [Hierarchical Clustering](#) (page 51)
- [K-Means Clustering](#) (page 54)
- [Marker Selection](#) (page 57)

### Standard Heatmap

A standard heatmap is a visualization of biomarker data points with no indication of patterns, groupings, or differentiation among the data points.

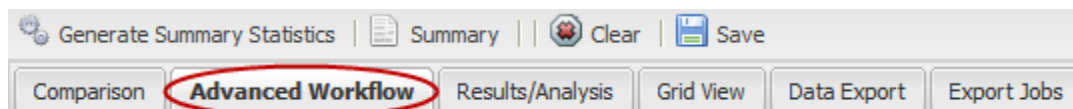
#### To generate a standard heatmap:

1. Run tranSMART, then click the **Dataset Explorer** tab.
2. Define the cohorts you wish to analyze by dragging one or more concepts from a study into empty subset definition boxes. For more information, see [Populating the Study Groups](#) on page 35.

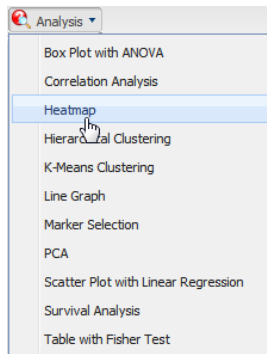


To compare two subsets, you may drag an additional concept into the Subset 2 comparison box.

3. Click the **Advanced Workflow** tab:

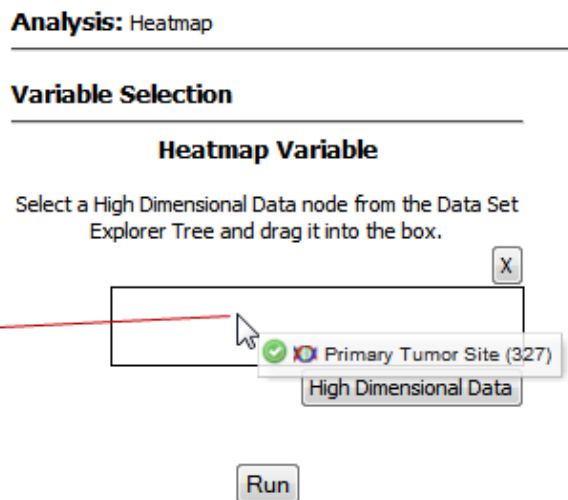


4. Select **Heatmap** from the **Analysis** dropdown menu:



The Variable Selection section appears.

5. Define the heatmap variable by selecting a high dimensional data node from the Dataset Explorer tree and dragging it into the Heatmap Variable definition box:



**i** High dimensional data nodes are indicated by the icon (X D) to the left of study data.

6. Click the **High Dimensional Data** button.

The Compare Subsets-Pathway Selection dialog appears.

7. Specify the platform and other factors of interest.

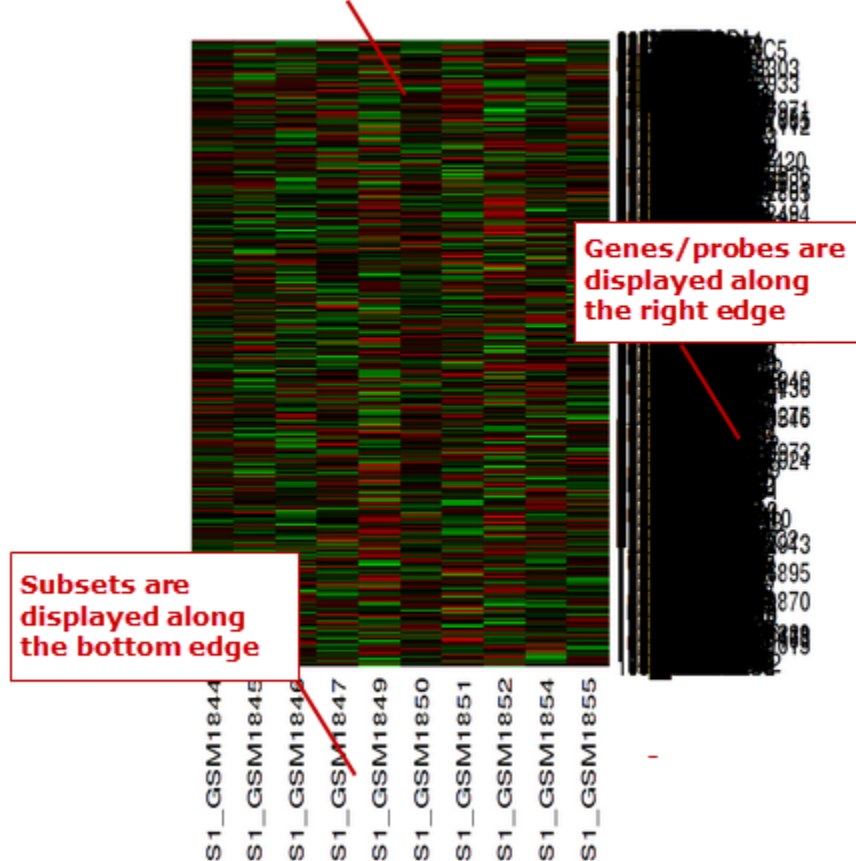
For more information, see [High Dimensional Data](#) on page 79.

8. Click **Apply Selections**.

9. Click **Run**.

Your analysis appears below:

A heatmap is a display of differential expression. Individual values contained in the matrix are represented by colors. Higher-than-normal expression is displayed in red. Lower-than-normal expression is displayed in green.



## Hierarchical Clustering

Hierarchical clustering is a visualization of patterns of related data points in gene expression data.

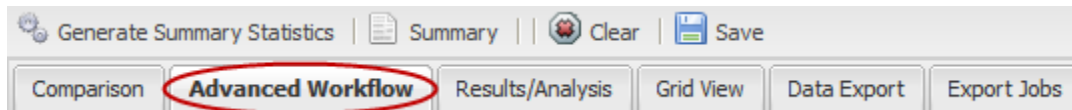
**To generate a hierarchical clustering heatmap:**

1. Run tranSMART, then click the **Dataset Explorer** tab.
2. Define the cohorts you wish to analyze by dragging one or more concepts from a study into empty subset definition boxes. For more information, see [Populating the Study Groups](#) on page 35.

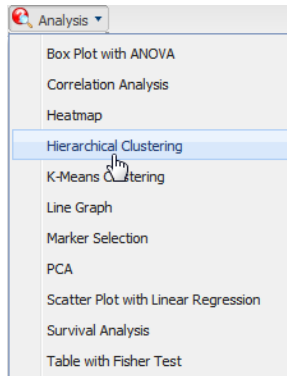


To compare two subsets, you may drag an additional concept into the Subset 2 comparison box.

- Click the **Advanced Workflow** tab:

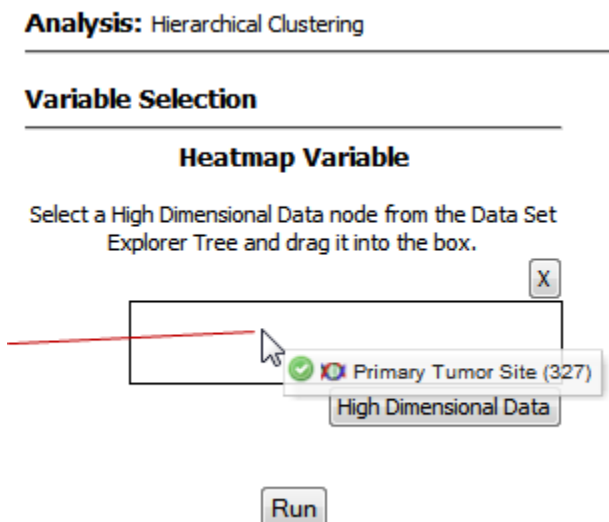


- Select **Hierarchical Clustering** from the **Analysis** dropdown menu:



The Variable Selection section appears.

- Define the heatmap variable by selecting a high dimensional data node from the Dataset Explorer tree and dragging it into the Heatmap Variable definition box:



High dimensional data nodes are indicated by the icon (📊) to the left of study data.

- Click the **High Dimensional Data** button.

The Compare Subsets-Pathway Selection dialog appears.

- Specify the platform and other factors of interest.

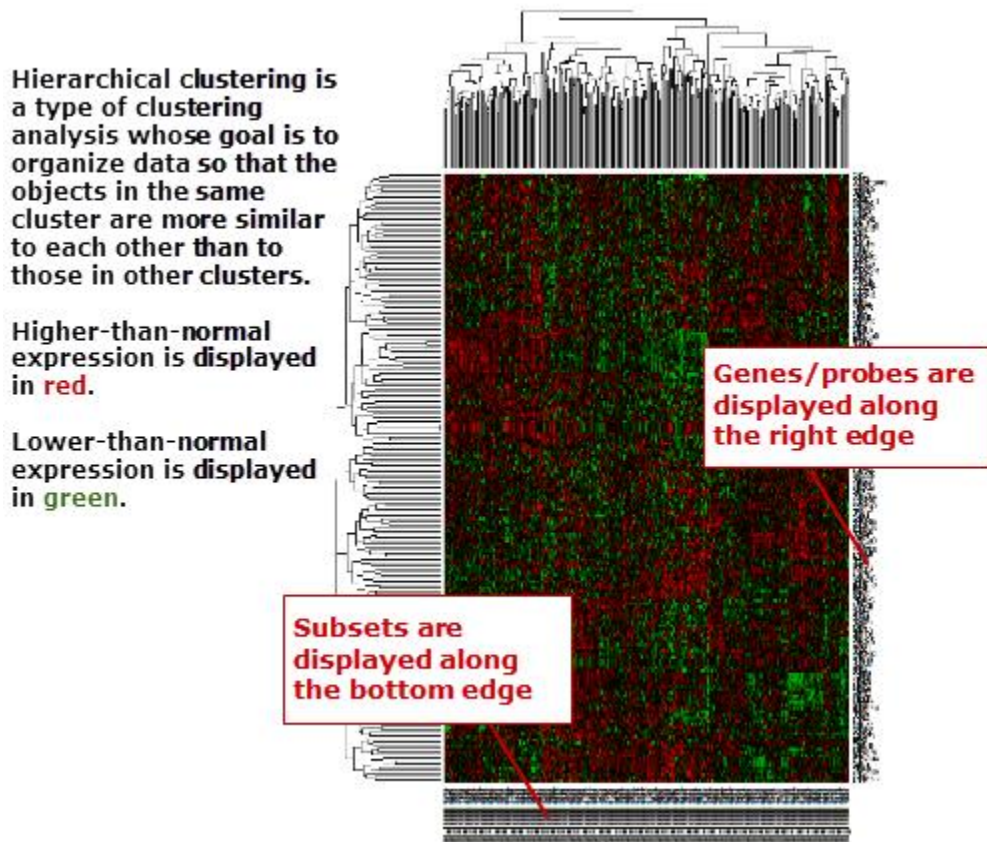
For more information, see [High Dimensional Data](#) on page 79.

- Click **Apply Selections**.



9. Click **Run**.

Your analysis appears below:



To read more about Hierarchical Clustering, visit:  
<http://www.ics.uci.edu/~epstein/280/cluster.html>

## K-Means Clustering

K-Means clustering is a visualization of groupings of the most closely related data points, based on the number of groupings you specify.



The K-Means analysis clusters columns together – rows are not clustered.

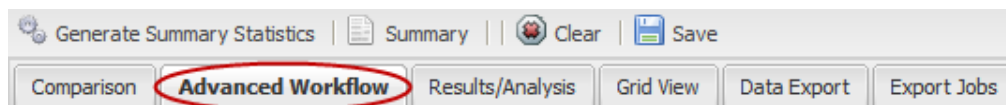
### To generate a k-means clustering heatmap:

1. Run tranSMART, then click the **Dataset Explorer** tab.
2. Define the cohorts you wish to analyze by dragging one or more concepts from a study into empty subset definition boxes. For more information, see [Populating the Study Groups](#) on page 35.

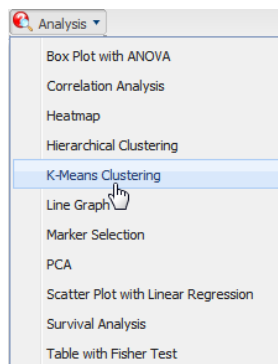


To compare two subsets, you may drag an additional concept into the Subset 2 comparison box.

3. Click the **Advanced Workflow** tab:



4. Select **K-Means Clustering** from the **Analysis** dropdown menu:



The Variable Selection section appears.

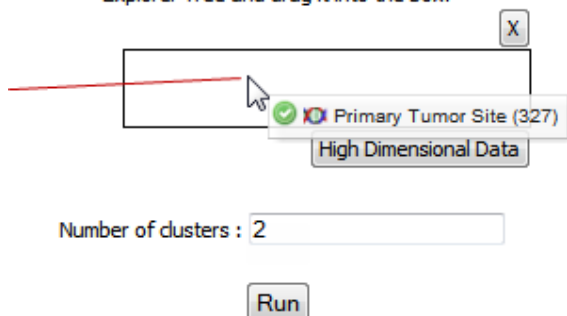
5. Define the heatmap variable by selecting a high dimensional data node from the Dataset Explorer tree and dragging it into the Heatmap Variable definition box:

**Analysis:** K-Means Clustering

### Variable Selection

#### Heatmap Variable

Select a High Dimensional Data node from the Data Set Explorer Tree and drag it into the box.



Number of clusters : 2

Run



High dimensional data nodes are indicated by the icon (X) to the left of study data.

6. Click the **High Dimensional Data** button.

The Compare Subsets-Pathway Selection dialog appears.

7. Specify the platform and other factors of interest.

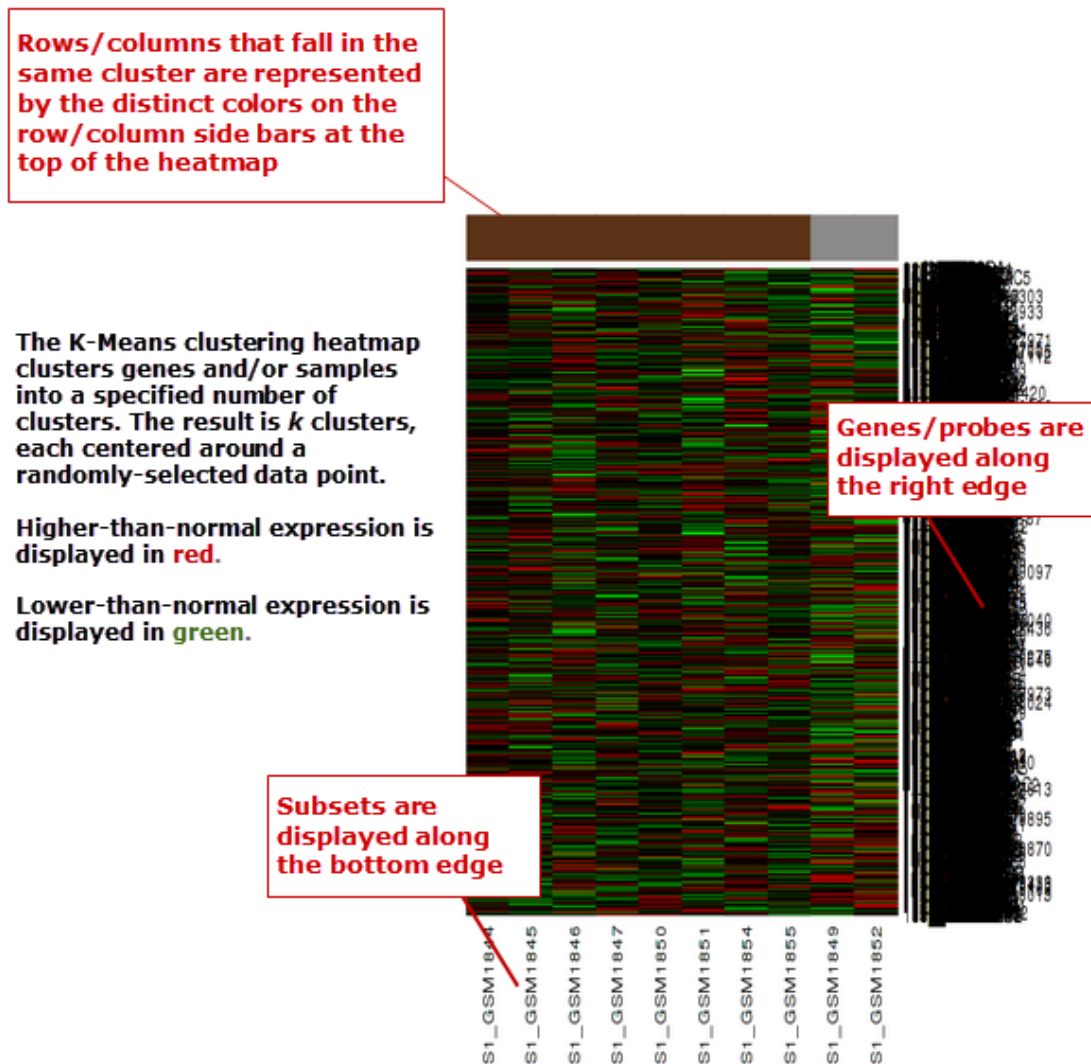
For more information, see [High Dimensional Data](#) on page 79.

8. Click **Apply Selections**.

9. In the **Number of clusters** field, type a numerical value.

10. Click **Run**.

Your analysis appears below:



To read more about K-Means Clustering, visit:  
<http://www.ics.uci.edu/~eppstein/280/cluster.html>

## Marker Selection

A marker selection heatmap is a visualization of differentially expressed genes in distinct phenotypes. Specifically, the algorithm determines the set of genes which is most differentially expressed between the two subsets. This list of differentially expressed genes is subsequently presented in a table, along with a variety of accompanying statistics.

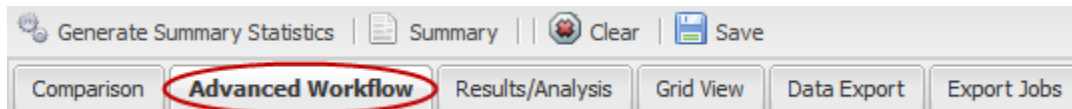
### To generate a marker selection heatmap:

1. Run tranSMART, then click the **Dataset Explorer** tab.
2. Define the cohorts you wish to analyze by dragging one or more concepts from a study into empty subset definition boxes. For more information, see [Populating the Study Groups](#) on page 35.

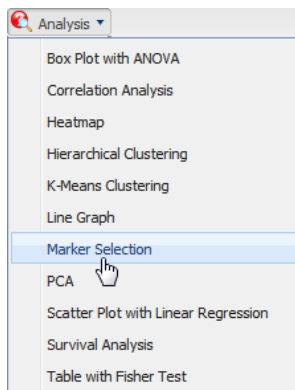


Two subsets must be specified when using a Marker Selection heatmap.

3. Click the **Advanced Workflow** tab:



4. Select **Marker Selection** from the **Analysis** dropdown menu:



The Variable Selection section appears.

- Define the required variable by selecting a high dimensional data node from the Dataset Explorer tree and dragging it into the Marker Variable definition box:

**Analysis:** Marker Selection

---

**Variable Selection**

---

**Marker Variable**

Select a High Dimensional Data node from the Data Set Explorer Tree and drag it into the box.

✔

 Primary Tumor Site (327)  
High Dimensional Data

Number of Markers :

High dimensional data nodes are indicated by the icon () to the left of study data.

- Click the **High Dimensional Data** button.

The Compare Subsets-Pathway Selection dialog appears.

- Specify the platform and other factors of interest.

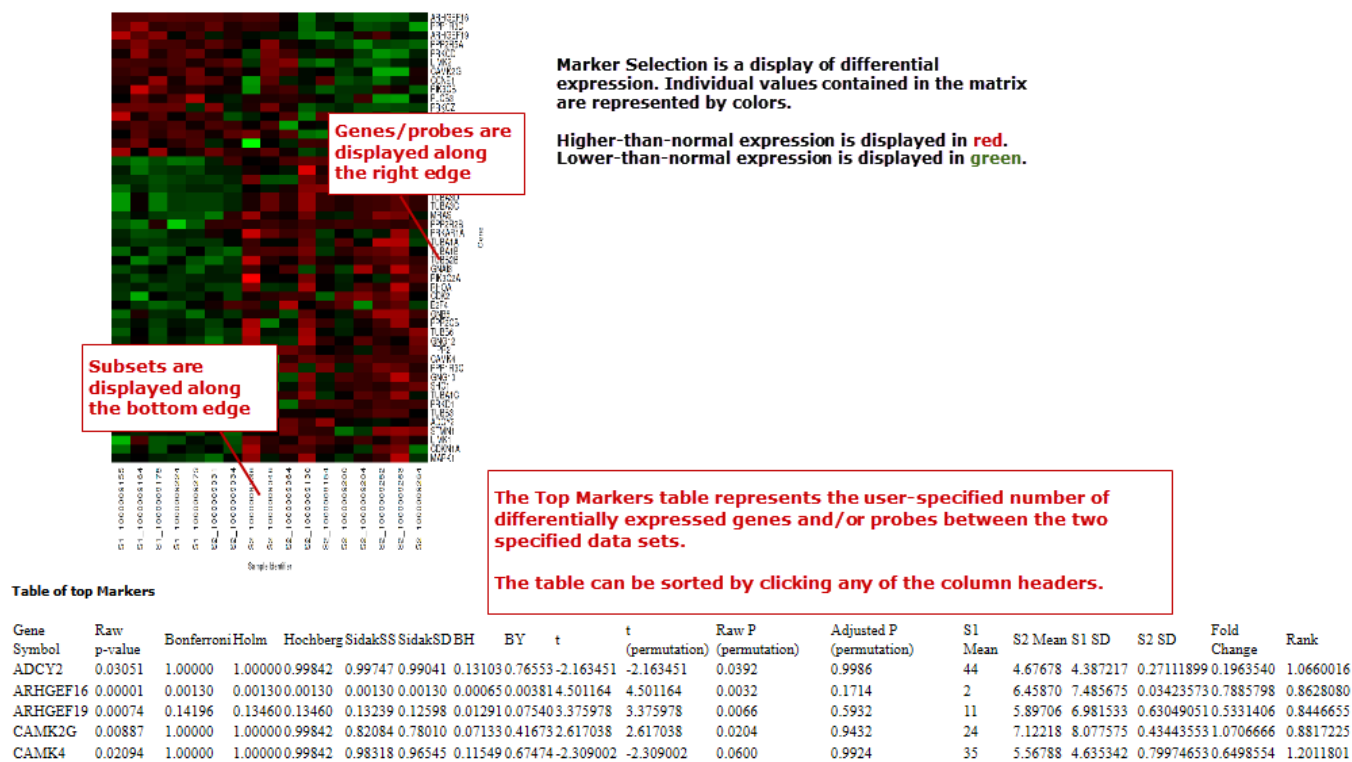
For more information, see [High Dimensional Data](#) on page 79.

- Click **Apply Selections**.

In the **Number of Markers** field, type a numerical value. This will determine the number of differentially expressed genes that are returned.

9. Click **Run**.

Your analysis appears below:



For more information on the analyses used in Marker Selection, visit:  
<http://mathworld.wolfram.com/bonferronicorrection.html>

## Generating Advanced Analyses

Advanced analyses include:

- [Box Plot with ANOVA](#) (page 60)
- [Correlation Analysis](#) (page 62)
- [Line Graph Analysis](#) (page 64)
- [Principal Component Analysis](#) (page 65)
- [Scatter Plot with Linear Regression](#) (page 68)
- [Survival Analysis](#) (page 70)
- [Table with Fisher Test Analysis](#) (page 73)

## Box Plot with ANOVA

A box plot with ANOVA analysis displays a box and whisker plot with corresponding analysis of variance in the sample(s).

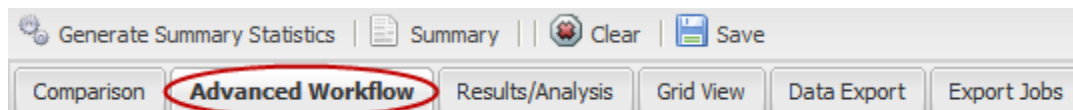
### To perform a box plot with ANOVA analysis:

1. Run tranSMART, then click the **Dataset Explorer** tab.
2. Define the cohorts you wish to analyze by dragging one or more concepts from a study into empty subset definition boxes. For more information, see [Populating the Study Groups](#) on page 35.

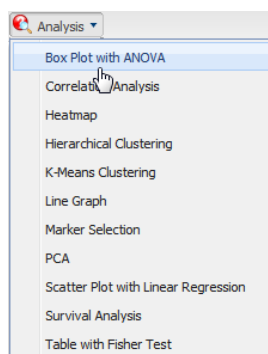


Only one subset may be specified in this analysis. Information in Subset 2 will be ignored.

3. Click the **Advanced Workflow** tab:



4. Select **Box Plot with ANOVA** from the **Analysis** dropdown menu:



The Variable Selection section appears. You will need to define what variables in the study are independent, and what variables are dependent. At least one of the variables should be continuous (for example, Age), and one should be a categorical value (for example, Tissue Type).



If the *independent variable* defines the groups, boxes will be plotted horizontally. If the *dependent variable* defines the groups, boxes will be plotted vertically.

5. Define the variables.

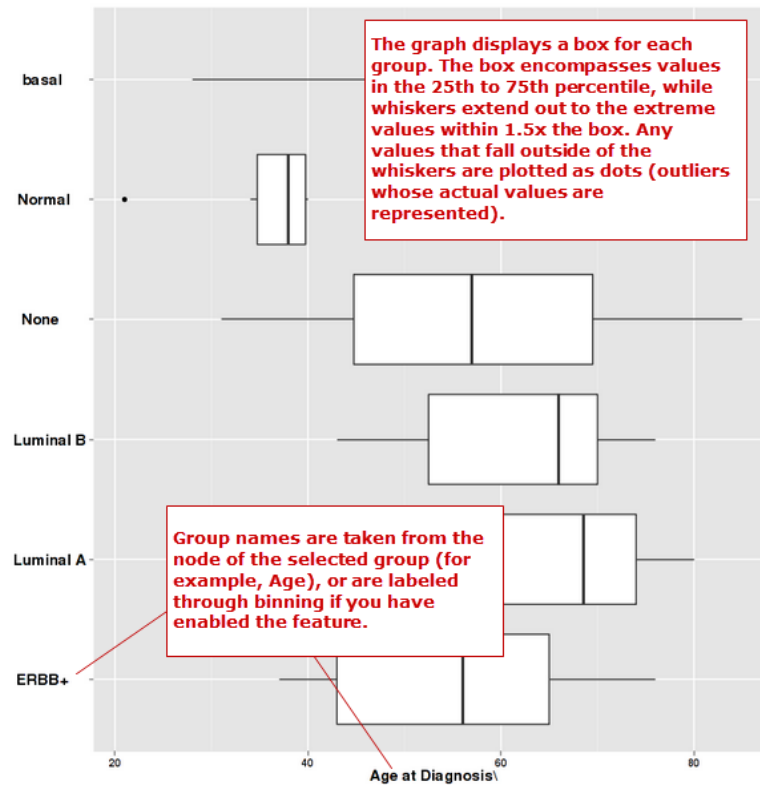


In this example, the data binning feature is not used. For future reference, data binning refers to a pre-processing technique used to reduce minor observation errors. Clusters of data are replaced by a value representative of that cluster (the central value). For information on binning, see [Data Binning](#) on page 74.



6. Click **Run**.

Your analysis appears below:

**ANOVA Result**

|         |         |
|---------|---------|
| p-value | 0.00181 |
| F value | 4.11    |

F value and p-value produced by ANOVA calculations on all groups.

| Group     | Mean | n  |
|-----------|------|----|
| ERBB+     | 54.5 | 11 |
| Luminal A | 64.3 | 28 |
| Luminal B | 62.1 | 11 |
| None      | 57.5 | 44 |
| Normal    | 37.3 | 6  |

The mean and population size of each group.

**Pairwise t-Test p-Values**

|           | ERBB+    | Luminal A | Luminal B | None     | Normal |
|-----------|----------|-----------|-----------|----------|--------|
| Luminal A | 5.19e-02 | NA        | NA        | NA       | NA     |
| Luminal B | 2.07e-01 | 6.59e-01  | NA        | NA       | NA     |
| None      | 5.34e-01 | 4.56e-02  | 3.28e-01  | NA       | NA     |
| Normal    | 1.65e-02 | 3.65e-05  | 6.65e-04  | 1.20e-03 | NA     |

Illustrates the p-value result after t-tests have been performed on each pair of groups.

## Correlation Analysis

In a correlation analysis, you are using statistical correlation to assess the relationship between variables.

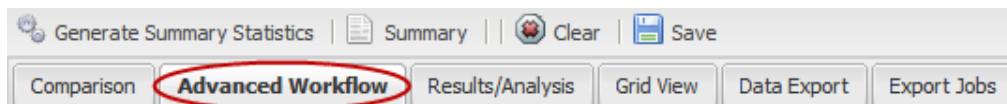
### To perform a correlation analysis:

1. Run tranSMART, then click the **Dataset Explorer** tab.
2. Define the cohorts you wish to analyze by dragging one or more concepts from a study into empty subset definition boxes. For more information, see [Populating the Study Groups](#) on page 35.

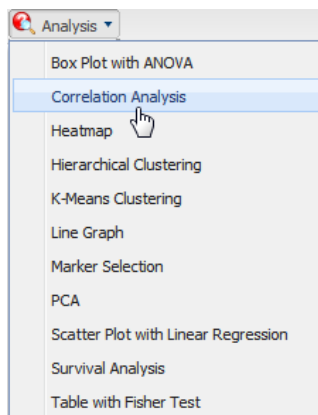


Only one subset may be specified in this analysis. Information in Subset 2 will be ignored.

3. Click the **Advanced Workflow** tab:

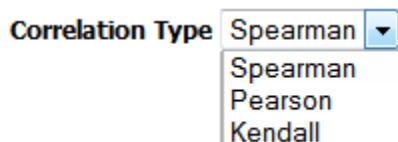


4. Select **Correlation Analysis** from the **Analysis** dropdown menu:



The Variable Selection section appears. You will need to define two or more continuous variables (for example, Age).

5. Define the variables.
6. Under **Correlation Type**, select the analysis you wish to perform from the dropdown menu:



The analyses listed under **Correlation Type** refer to different regression algorithms.

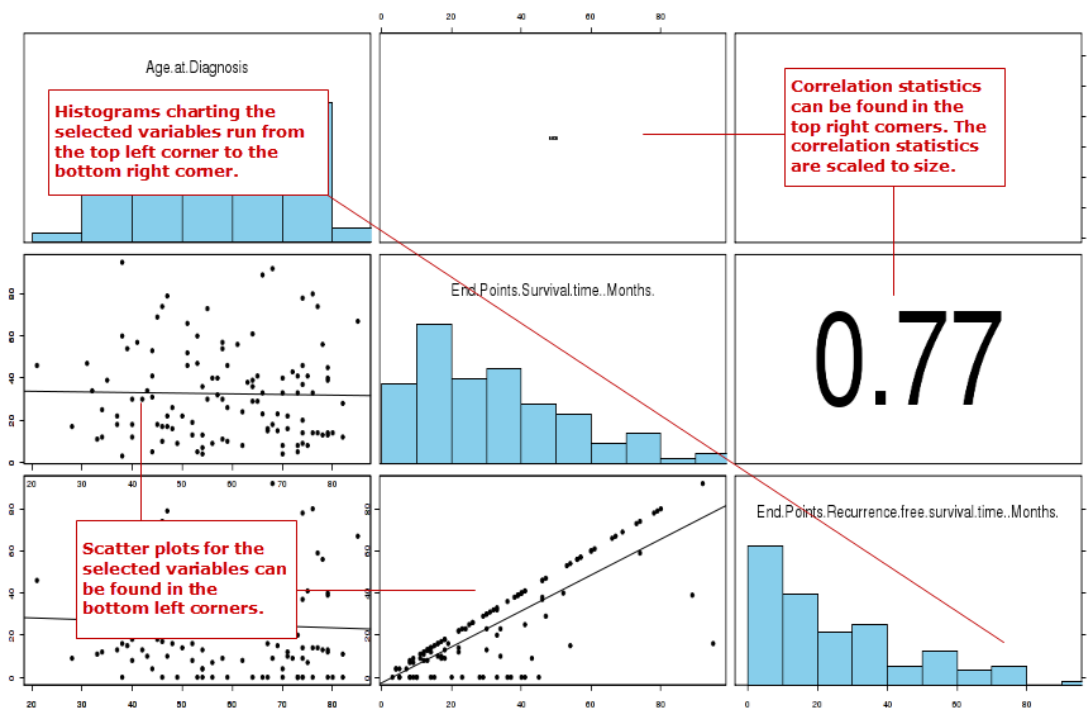
7. Click **Run**.

Your analysis appears below:

Correlation Table (p-values on top right half, correlation coefficient on bottom left)

|   | Age at Diagnosis | End Points Survival time Months | End Points Recurrence free survival time Months |
|---|------------------|---------------------------------|---|
| Age at Diagnosis                                | 1.000000         | 0.704676                        | 0.230513  |
| End Points Survival time Months                 | -0.035722        | 1.000000                        | 0.000000  |
| End Points Recurrence free survival time Months | -0.112685        | 0.766856                        | 1.000000  |

The correlation table displays statistical analyses of the selected variables.



## Line Graph Analysis

In a line graph analysis, you are comparing time or measurement concepts with a subset of subjects in a study (for example, treatment groups).



Each group will be plotted as a distinct line on the graph.

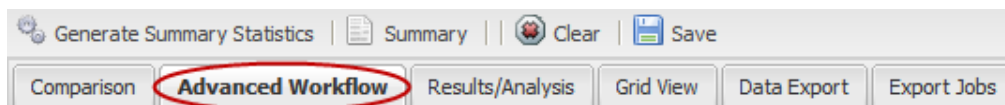
### To perform a line graph analysis:

1. Run tranSMART, then click the **Dataset Explorer** tab.
2. Define the cohorts you wish to analyze by dragging one or more concepts from a study into empty subset definition boxes. For more information, see [Populating the Study Groups](#) on page 35.

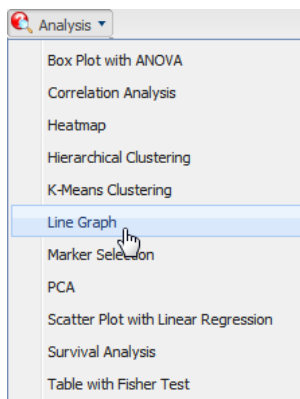


Only one subset may be specified in this analysis. Information in Subset 2 will be ignored.

3. Click the **Advanced Workflow** tab:



4. Select **Correlation Analysis** from the **Analysis** dropdown menu:

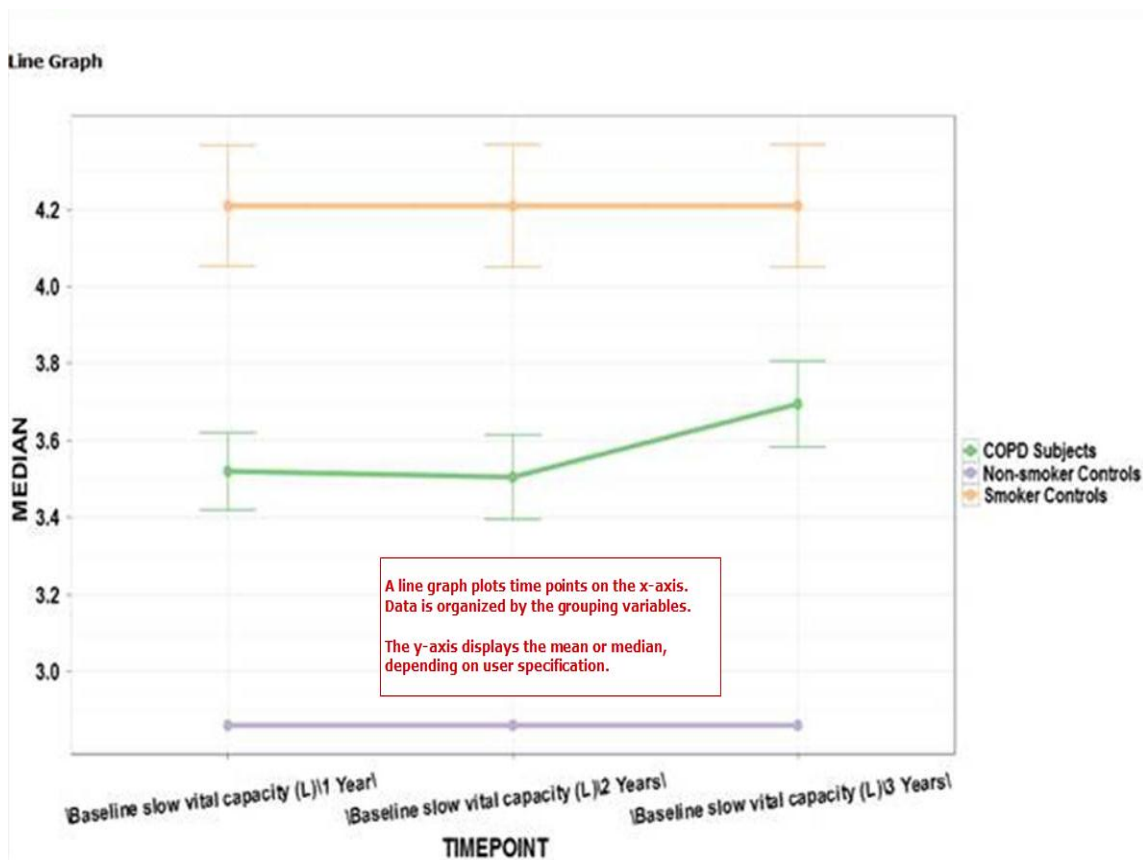


The Variable Selection section appears. You will need to define time or measurement concept(s), and one or more grouping concepts (for example, treatment groups).

5. Define the variables.
6. Under **Graph Type**, select the type of line graph you wish to be displayed.

7. Click **Run**.

Your analysis appears below:



## Principal Component Analysis

In a principal component analysis (PCA), the total number of variables in the dataset is reduced to a smaller number of variables – the principle components of the dataset.

Principal component variables are calculated from correlated variables in the total dataset. In other words, the principal component analysis is a workflow used to identify variance in a dataset. The analysis can be run on an entire microarray chip, or on a pathway.

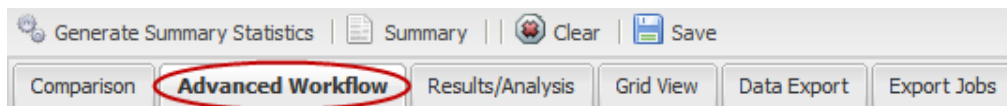
### To perform a principal component analysis:

1. Run tranSMART, then click the **Dataset Explorer** tab.
2. Define the cohorts you wish to analyze by dragging one or more concepts from a study into empty subset definition boxes. For more information, see [Populating the Study Groups](#) on page 35.

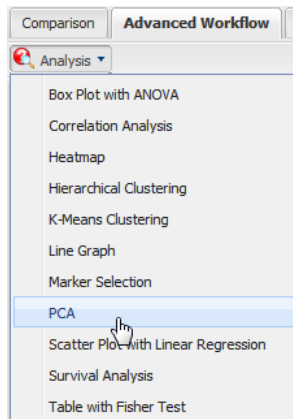


Only one subset may be specified in this analysis. Information in Subset 2 will be ignored.

- Click the **Advanced Workflow** tab:

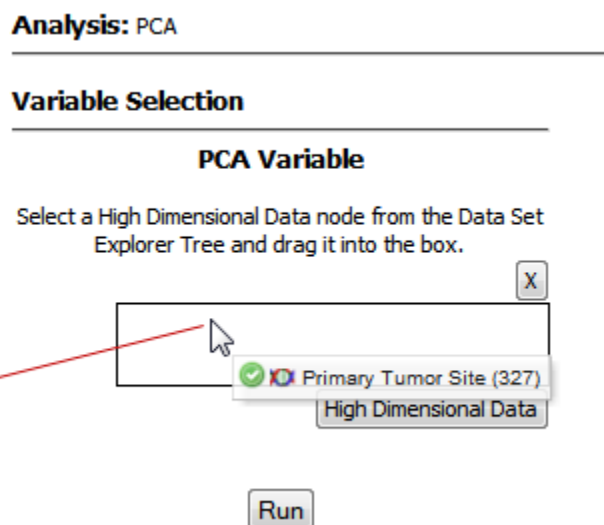



- Select **PCA** from the **Analysis** dropdown menu:



The Variable Selection section appears.

- Define the heatmap variable by selecting a high dimensional data node from the Dataset Explorer tree and dragging it into the PCA Variable definition box:



High dimensional data nodes are indicated by the (  ) icon to the left of study data.

- Click the **High Dimensional Data** button.

The Compare Subsets-Pathway Selection dialog appears.

- Specify the platform and other factors of interest.

For more information, see [High Dimensional Data](#) on page 79.

8. Click **Apply Selections**.

9. Click **Run**.

Your analysis appears below:

Component Summary

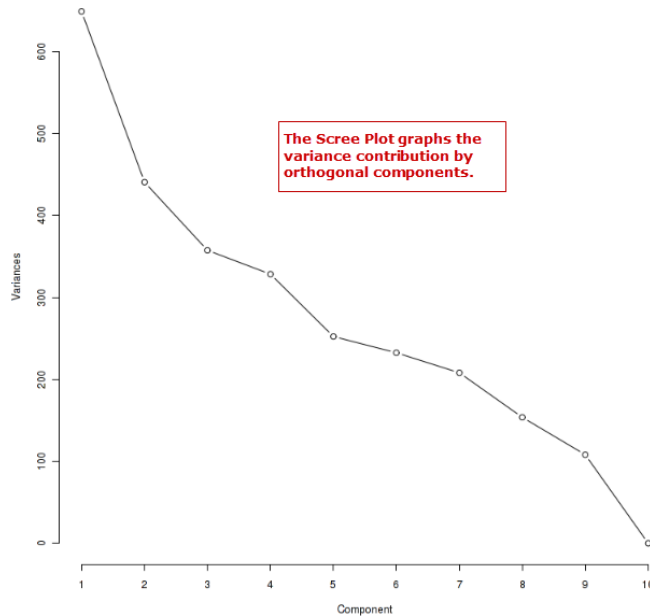
| Primary Component | Eigen Value | Percent Variance |
|-------------------|-------------|------------------|
| PC1               | 649.09441   | 23.77064         |
| PC2               | 440.686     | 16.13847         |
| PC3               | 357.5004    | 13.09211         |
| PC4               | 328.47809   | 12.02927         |
| PC5               | 252.33128   | 9.24068          |
| PC6               | 232.52903   | 8.5155           |
| PC7               | 208.12087   | 7.62164          |
| PC8               | 153.7603    | 5.63089          |
| PC9               | 108.15558   | 3.96079          |
| PC10              | 0           | 0                |

A Principal Component Analysis (PCA) is commonly used as a tool in exploratory data analysis.

Data is split into orthogonal components, and the genes/probes that contribute the most to the components are displayed.

The Component Summary table displays the orthogonal components that your data has been broken into, and how much of the overall variance they are contributing to the variance in the total data set (percent variance).

Scree Plot



Gene list by proximity to Component

| Component 1          | Component 2        | Component 3        | Component 4          |
|----------------------|--------------------|--------------------|----------------------|
| X5404_OGN -0.061     | X4728_ANXA8L1 0.06 | X185_MKI67 -0.067  | X569_YME1L1 -0.068   |
| X1228_OPA1 -0.055    | X2045_SORBS2 0.058 | X4398_TGFA 0.067   | X4278_PPP1R14C 0.067 |
| X2129_DENND4B -0.054 | X1094_DST          |                    | X3571_CNTN3 0.062    |
| X738_ILIRA2 -0.053   | X1130_MGST1        |                    | X2054_MS4A6A 0.061   |
| X2573_FABP7 0.052    | X2787_KRT17 0.056  | X2481_SORL1 0.059  | X4996_CSMD1 -0.06    |
| X4195_CHI3L2 0.052   | X4340_MMP7 0.055   | X4967_CRABP1 0.059 | X4689_ACS3 -0.056    |

The Gene List table lists genes that make up each orthogonal component.



For more information regarding PCAs, see: <http://psb.stanford.edu/psb-online/proceedings/psb00/raychaudhuri.pdf>.

## Scatter Plot with Linear Regression

A scatter plot displays values for two variables within a dataset, with a line that best fits the slope of the data.

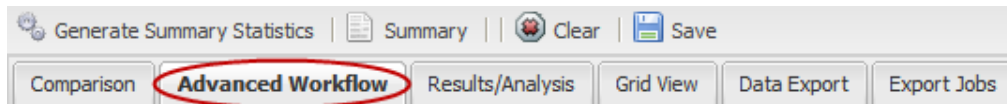
### To perform a scatter plot with linear regression analysis:

1. Run tranSMART, then click the **Dataset Explorer** tab.
2. Define the cohorts you wish to analyze by dragging one or more concepts from a study into empty subset definition boxes. For more information, see [Populating the Study Groups](#) on page 35.

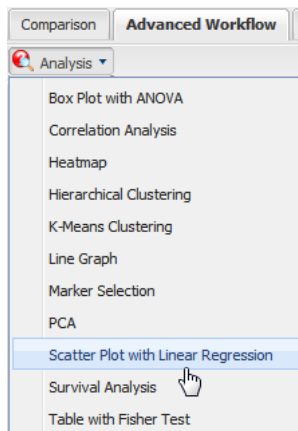


Only one subset may be specified in this analysis. Information in Subset 2 will be ignored.

3. Click the **Advanced Workflow** tab above Subset 1:



4. Select **Scatter Plot with Linear Regression** from the **Analysis** dropdown menu:



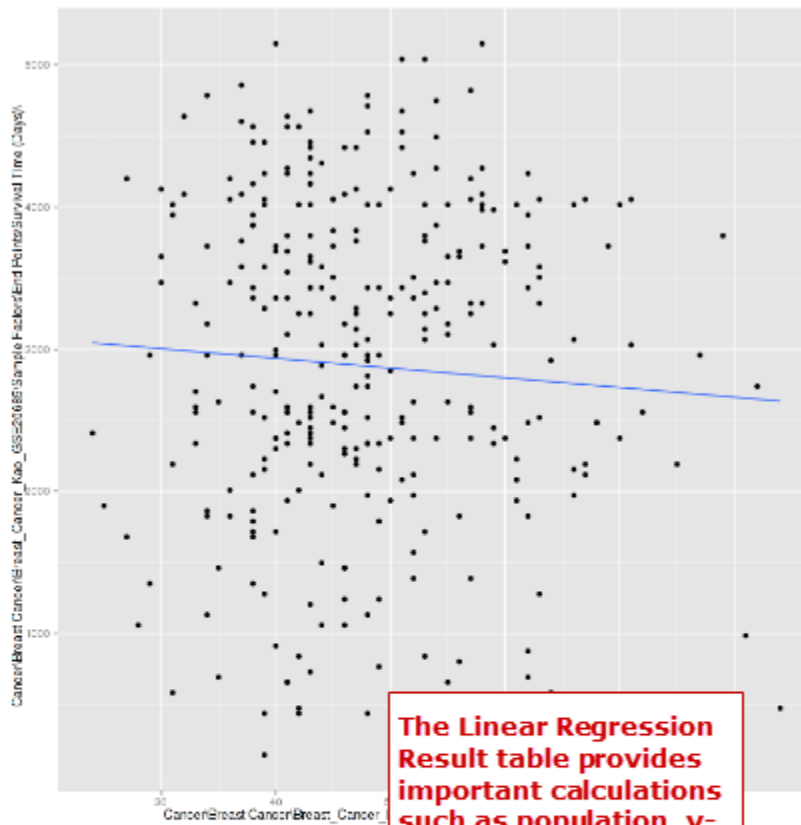
The Variable Selection section appears. You will need to define what variables in the study are independent, and what variables are dependent. Both variables should be continuous (for example, Age).

5. Define the variables.



6. Click **Run**.

Your analysis appears below:

**Scatter Plot****Linear Regression Result**

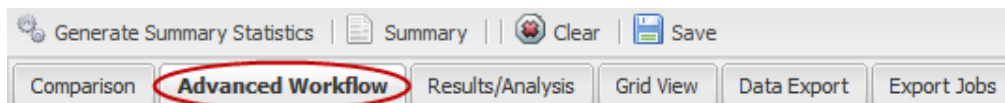
|                    |          |
|--------------------|----------|
| Number of Subjects | 327      |
| Intercept          | 3210     |
| Slope              | -6.83    |
| r-squared          | 0.00393  |
| adjusted r-squared | 0.000868 |
| p-value            | 0.258    |

## Survival Analysis

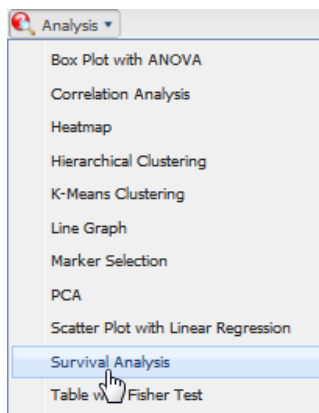
A survival analysis displays time-to-event data.

### To perform a survival analysis:

1. Run tranSMART, then click the **Dataset Explorer** tab.
2. Define the cohorts you wish to analyze by dragging one or more concepts from a study into empty subset definition boxes. For more information, see [Populating the Study Groups](#) on page 35.
3. Click the **Advanced Workflow** tab above Subset 1:



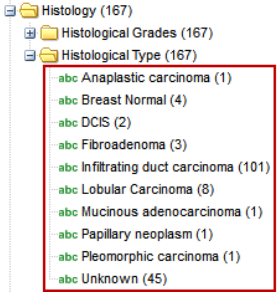
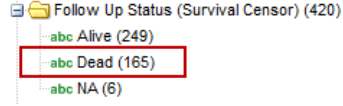
4. Select **Survival Analysis** from the **Analysis** dropdown menu:



The Variable Selection section appears.

5. Define the following variables:

| Variable | Required? | Definition                        | Example                           |
|----------|-----------|-----------------------------------|-----------------------------------|
| Time     | Yes       | A numeric field within tranSMART. | Survival at Follow Up (Years)<br> |

| Variable        | Required? | Definition   | Example   |
|-----------------|-----------|--|---|
| Category        | No        | <p>A concept that is dragged into this input will dictate the groups into which the data will be split in order to compare their survival times.</p> <p>If this variable is continuous, it requires binning. For details, see <a href="#">Data Binning Using Survival Analysis</a> on page 76.</p> | <p>Cancer Stage</p>  |
| Censoring Value | No        | <p>Specifies which patients had the event whose time is being measured. For example, if the Time variable selected is <b>Overall Survival Time (Years)</b>, an appropriate censoring variable is <b>Patient Death</b>.</p>   | <p>Dead</p>          |

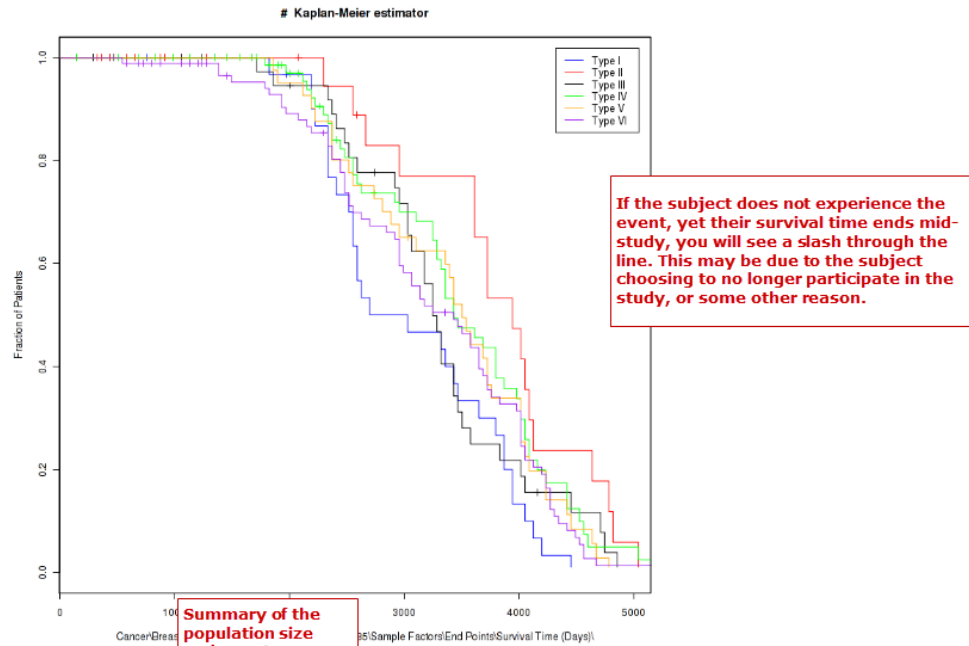


In this example, the data binning feature is not used. For future reference, data binning refers to a pre-processing technique used to reduce minor observation errors. Clusters of data are replaced by a value representative of that cluster (the central value). For information on data binning, see [Data Binning Using Survival Analysis](#) on page 76.

6. Click **Run**.

Your analysis appears below:

Survival Curve



Cox Regression Result

|                    |     |
|--------------------|-----|
| Number of Subjects | 327 |
| Number of Events   | 244 |

Results of Cox Regression  
(proportional hazards model)

| Subset   | Cox Coefficient | Hazards Ratio | Lower Range of Hazards Ratio, 95% Confidence Interval | Upper Range of Hazards Ratio, 95% Confidence Interval |
|----------|-----------------|---------------|---|---|
| Type II  | -0.8941         | 0.4090        | 0.2231  | 0.7497  |
| Type III | -0.3775         | 0.6855        | 0.4143  | 1.1343  |
| Type IV  | -0.5474         | 0.5785        | 0.3675  | 0.9105  |
| Type V   | -0.4294         | 0.6509        | 0.4021  | 1.0536  |
| Type VI  | -0.3595         | 0.6980        | 0.4556  | 1.0803  |

Summary of the survival  
curve fit (by bin if enabled)

Survival Curve Fitting Summary

| Subset   | Number of Subjects | Max Subjects | Subjects at Start | Number of Events | Median Time Value | Lower Range of Time Variable, 95% Confidence Interval | Upper Range of Time Variable, 95% Confidence Interval |
|----------|--------------------|--------------|-------------------|------------------|-------------------|---|---|
| Type I   | 37                 | 37           | 37                | 30               | 3032              | 2557  | 3799  |
| Type II  | 34                 | 34           | 34                | 17               | 3945              | 3616  | 4639  |
| Type III | 41                 | 41           | 41                | 32               | 3251              | 3068  | 3506  |
| Type IV  | 81                 | 81           | 81                | 52               | 3433              | 3287  | 3981  |
| Type V   | 41                 | 41           | 41                | 38               | 3506              | 3105  | 4018  |
| Type VI  | 93                 | 93           | 93                | 75               | 3423              | 2959  | 3726  |

## Table with Fisher Test Analysis

A Fisher Test analysis examines the significance of associated variables.

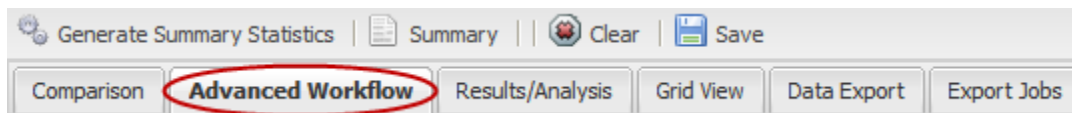
### To perform a table with fisher test analysis:

1. Run tranSMART, then click the **Dataset Explorer** tab.
2. Define the cohorts you wish to analyze by dragging one or more concepts from a study into empty subset definition boxes. For more information, see [Populating the Study Groups](#) on page 35.

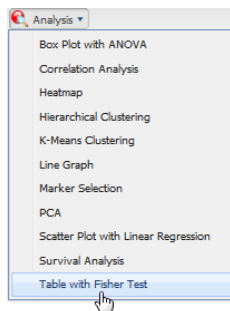


Only one subset may be specified in this analysis. Information in Subset 2 will be ignored.

3. Click the **Advanced Workflow** tab:



4. Select **Table with Fisher Test** from the **Analysis** dropdown menu:



The Variable Selection section appears. You will need to define what variables in the study are independent, and what variables are dependent. Both variables should be categorical values (for example, Tissue Type).



In this example, the data binning feature is not used. For future reference, data binning refers to a pre-processing technique used to reduce minor observation errors. Clusters of data are replaced by a value representative of that cluster (the central value). For information on data binning, see [Data Binning](#) on page 74.

5. Define the variables.

6. Click **Run**.

Your analysis appear below:

|              |               |             |
|--------------|---------------|-------------|
|              | <b>Female</b> | <b>Male</b> |
| <b>Alive</b> | 105           | 135         |
| <b>Dead</b>  | 67            | 89          |

**Population of each category's members.**

|                            |          |
|----------------------------|----------|
| <b>Fisher test p-value</b> | =0.918   |
| $\chi^2$                   | =0.00286 |
| $\chi^2$ p-value           | =0.957   |

**Chi-squared distribution values.**

## Data Binning

Data binning refers to a pre-processing technique used to reduce observation errors and to allow continuous variables to become categorical. Clusters of data are replaced by a value representative of that cluster (the central value).



The data displayed after binning represents the data available in the study. If, for example, you have selected to bin based on date range (0-10 years of age), yet there is only data available for subjects eight years old and up, the bin will display the age range as 8-10.

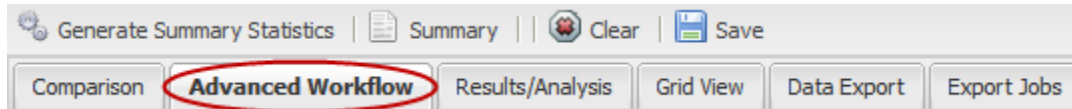
### Data Binning Using Box Plot with ANOVA

When conducting a Box Plot with ANOVA analysis, at least one of the variables selected should be a continuous variable (for example, age), and the other should be a categorical value (for example, tumor stage).

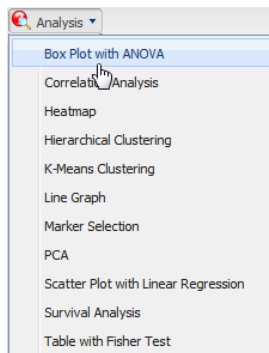
A continuous variable can be viewed as a categorical value using the binning feature, described below. Alternatively, binning can also be used to categorize data. For example, if histological grade with values such as Well Defined, Moderately Well Defined, and Poorly Defined are selected, you can group Moderately Well Defined with Poorly Defined and treat them as one group for the purposes of this analysis.

**To use the data binning feature with a box plot analysis:**

1. Run tranSMART, then click the **Dataset Explorer** tab.
2. Define the cohorts you wish to analyze by dragging one or more concepts from a study into empty subset definition boxes. For more information, see [Populating the Study Groups](#) on page 35.
3. Click the **Advanced Workflow** tab:



4. Select **Box Plot with ANOVA** from the **Analysis** dropdown menu:



The Variable Selection section appears. You will need to define what variables in the study are independent, and what variables are dependent. At least one of the variables should be continuous (for example, Age), and one should be a categorical value (for example, Tissue Type).

5. Define the variables.
6. Under **Binning**, click **Enable**:



7. Define the following:

| Field         | Description   | Comments   |
|---------------|---|--|
| Variable      | Select which variable should define the groups (Independent or Dependent) from the dropdown menu.       | If the <i>independent variable</i> defines the groups, boxes will be plotted horizontally. If the <i>dependent variable</i> defines the groups, boxes will be plotted vertically |
| Variable Type | Select whether the variable you have defined above is continuous or categorical from the dropdown menu. | A continuous variable can be turned into a categorical variable when you use the binning feature.  |

| Field                              | Description  | Comments   |          |       |       |   |       |   |       |   |       |   |                   |              |                                    |                      |  |              |  |                      |
|------------------------------------|--|--|----------|-------|-------|---|-------|---|-------|---|-------|---|-------------------|--------------|------------------------------------|----------------------|--|--------------|--|----------------------|
| Number of Bins                     | Type the number of bins you would like data to be organized in.  | This step may require trial and error based on how you wish to display data.   |          |       |       |   |       |   |       |   |       |   |                   |              |                                    |                      |  |              |  |                      |
| Bin Assignments                    | Select how you would like data to be binned from the dropdown menu.<br><b>Note:</b> This feature can only be used when the variable type selected above is continuous. | <ul style="list-style-type: none"><li>▪ <b>Evenly Distribute Population:</b> assigns bins based on the underlying data. For example, if the majority of the subjects in the study were elderly, bins based on age could look like: [(1-40), (40-80), (81-85), (86-90), (90-92)].</li><li>▪ <b>Evenly Spaced Bins:</b> creates bins based on the overall range of the variable. For example, if the majority of the subjects in the study were elderly, bins based on age could look like: [(1-20), (21-40), (41-60), (61-80), (81-100)].</li></ul>   |          |       |       |   |       |   |       |   |       |   |                   |              |                                    |                      |  |              |  |                      |
| Manual Binning                     | Select the checkbox if you wish to bin manually.<br><b>Note:</b> This is the only binning method available if you are attempting to bin a categorical variable type.   | Complete the binning form that populates as a result of checking the <b>Manual Binning</b> box.<br>For continuous data:<br><table border="1"><thead><tr><th>Bin Name</th><th>Range</th></tr></thead><tbody><tr><td>Bin 1</td><td><input type="text"/> - <input type="text"/></td></tr><tr><td>Bin 2</td><td><input type="text"/> - <input type="text"/></td></tr><tr><td>Bin 3</td><td><input type="text"/> - <input type="text"/></td></tr><tr><td>Bin 4</td><td><input type="text"/> - <input type="text"/></td></tr></tbody></table><br>For categorical data:<br><table><tr><td><b>Categories</b></td><td><b>Bin 1</b></td></tr><tr><td><div>&lt;- Drag To Bin -&gt;</div></td><td><input type="text"/></td></tr><tr><td></td><td><b>Bin 2</b></td></tr><tr><td></td><td><input type="text"/></td></tr></table> | Bin Name | Range | Bin 1 | <input type="text"/> - <input type="text"/> | Bin 2 | <input type="text"/> - <input type="text"/> | Bin 3 | <input type="text"/> - <input type="text"/> | Bin 4 | <input type="text"/> - <input type="text"/> | <b>Categories</b> | <b>Bin 1</b> | <div>&lt;- Drag To Bin -&gt;</div> | <input type="text"/> |  | <b>Bin 2</b> |  | <input type="text"/> |
| Bin Name                           | Range  |  |          |       |       |   |       |   |       |   |       |   |                   |              |                                    |                      |  |              |  |                      |
| Bin 1                              | <input type="text"/> - <input type="text"/>  |  |          |       |       |   |       |   |       |   |       |   |                   |              |                                    |                      |  |              |  |                      |
| Bin 2                              | <input type="text"/> - <input type="text"/>  |  |          |       |       |   |       |   |       |   |       |   |                   |              |                                    |                      |  |              |  |                      |
| Bin 3                              | <input type="text"/> - <input type="text"/>  |  |          |       |       |   |       |   |       |   |       |   |                   |              |                                    |                      |  |              |  |                      |
| Bin 4                              | <input type="text"/> - <input type="text"/>  |  |          |       |       |   |       |   |       |   |       |   |                   |              |                                    |                      |  |              |  |                      |
| <b>Categories</b>                  | <b>Bin 1</b>   |  |          |       |       |   |       |   |       |   |       |   |                   |              |                                    |                      |  |              |  |                      |
| <div>&lt;- Drag To Bin -&gt;</div> | <input type="text"/>   |  |          |       |       |   |       |   |       |   |       |   |                   |              |                                    |                      |  |              |  |                      |
|                                    | <b>Bin 2</b>   |  |          |       |       |   |       |   |       |   |       |   |                   |              |                                    |                      |  |              |  |                      |
|                                    | <input type="text"/>   |  |          |       |       |   |       |   |       |   |       |   |                   |              |                                    |                      |  |              |  |                      |

8. Click **Run**.

## Data Binning Using Survival Analysis

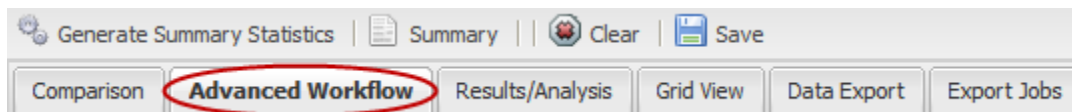
Data binning is used in survival analyses if the variable you wish to use is continuous (for example, age), but needs to be viewed as categorical data. Alternatively, it can be used to regroup categorical data. For example, if histological grade with values such as *Well Defined*, *Moderately Well Defined*, and *Poorly Defined* are selected, you can group *Moderately Well Defined* with *Poorly Defined* and treat them as one group for the purposes of this analysis.

### To use the data binning feature with a survival analysis:

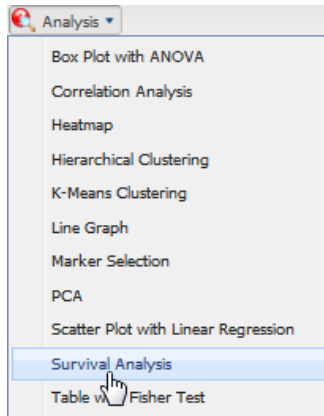
1. Run tranSMART, then click the **Dataset Explorer** tab.
2. Define the cohorts you wish to analyze by dragging one or more concepts from a study into empty subset definition boxes. For more information, see [Populating the Study Groups](#) on page 35.



3. Click the **Advanced Workflow** tab:



4. Select **Survival Analysis** from the **Analysis** dropdown menu:

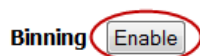


The Variable Selection section appears.

5. Define the variables:

| Variable           | Required? | Description   | Example           |
|--------------------|-----------|---|-------------------|
| Time               | Yes       | A time variable used in the study.  | Survival Time     |
| Category           | No        | A variable that you wish to use to sort the cohorts.<br><br>If the variable you wish to use is continuous (for example, age), the binning feature should be used. | Cancer Stage      |
| Censoring Variable | No        | A censoring variable (occurs when the value of a measurement/observation is partially known).   | Survival (Censor) |

6. Under **Binning**, click **Enable**:



## 7. Define the following:

| Field           | Description  | Comments  |          |       |       |   |       |   |       |   |       |   |
|-----------------|--|---|----------|-------|-------|---|-------|---|-------|---|-------|---|
| Variable Type   | Select whether the variable you have defined above is continuous or categorical.   | A continuous variable can be treated as a categorical variable when you use the binning feature.  |          |       |       |   |       |   |       |   |       |   |
| Number of Bins  | Type the number of bins you would like data to be organized in.  | This step may require trial and error based on how you wish to display data.  |          |       |       |   |       |   |       |   |       |   |
| Bin Assignments | Select how you would like data to be binned.<br><b>Note:</b> This feature can only be used when the variable type selected above is continuous.                      | <ul style="list-style-type: none"><li>Evenly Distribute Population: assigns bins based on the underlying data.<br/>For example, if the majority of the subjects in the study were elderly, bins based on age could look like: [(1-40), (40-80), (81-85), (86-90), (90-92)].</li><li>Evenly Spaced Bins: creates bins based on the overall range of the variable.<br/>For example, if the majority of the subjects in the study were elderly, bins based on age could look like: [(1-20), (21-40), (41-60), (61-80), (81-100)].</li></ul>                              |          |       |       |   |       |   |       |   |       |   |
| Manual Binning  | Select the checkbox if you wish to bin manually.<br><b>Note:</b> This is the only binning method available if you are attempting to bin a categorical variable type. | <p>Complete the binning form that populates as a result of checking the <b>Manual Binning</b> box.</p> <ul style="list-style-type: none"><li>For continuous data:<div><table><thead><tr><th>Bin Name</th><th>Range</th></tr></thead><tbody><tr><td>Bin 1</td><td>-</td></tr><tr><td>Bin 2</td><td>-</td></tr><tr><td>Bin 3</td><td>-</td></tr><tr><td>Bin 4</td><td>-</td></tr></tbody></table></div></li><li>For categorical data:<div><div><div>Categories</div><div>&lt;- Drag To Bin -&gt;</div></div><div><div>Bin 1</div><div>Bin 2</div></div></div></li></ul> | Bin Name | Range | Bin 1 | - | Bin 2 | - | Bin 3 | - | Bin 4 | - |
| Bin Name        | Range  |   |          |       |       |   |       |   |       |   |       |   |
| Bin 1           | -  |   |          |       |       |   |       |   |       |   |       |   |
| Bin 2           | -  |   |          |       |       |   |       |   |       |   |       |   |
| Bin 3           | -  |   |          |       |       |   |       |   |       |   |       |   |
| Bin 4           | -  |   |          |       |       |   |       |   |       |   |       |   |

8. Click **Run**.

## High Dimensional Data

The High Dimensional Data button available within the Advanced Workflow section of Dataset Explorer allows you to specify additional inputs for selected variables. These inputs help filter specific information of value (such as platforms, samples, or timepoints).

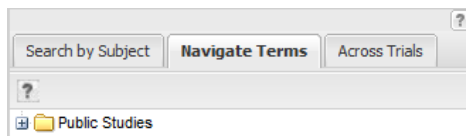


The High Dimensional Data feature must be used when you perform an analysis using high dimensional data (such as SNP, gene expression, RBM, etc.) symbolized by the DNA icon (🧬). Additionally, the High Dimensional Data feature cannot be used without high dimensional data.

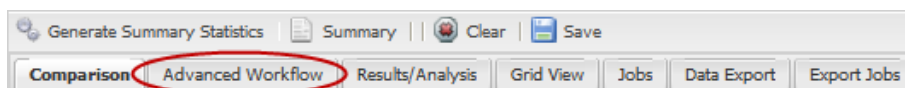
### To use the High Dimensional Data feature:

1. Click the tranSMART **Dataset Explorer** tab to display the Dataset Explorer window.
2. In the left pane of the Dataset Explorer window, click the **Navigate Terms** tab.

The navigation tree appears, showing the categories of available studies:



3. Select the study of interest.
4. Drag the study of interest into a subset definition box in Subset 1.
5. Click the **Advanced Workflow** tab above Subset 1:



6. Select the analysis you wish to perform, and define at least one variable using high dimensional data.
7. Click the **High Dimensional Data** button.

The Compare Subsets-Pathway Selection dialog appears.

8. Define the following available filters:



Dataset Explorer will attempt to pre-populate default values in the associated fields of the dialog based on the underlying data in the variable selection box.

| Filter                | Description  |
|-----------------------|--|
| Platform              | The platform type (for example, SNP, mRNA, etc.) used to collect biomarker data in the study.  |
| GPL Platform          | The specific name of the platform used in the study.   |
| Sample                | The type of sample tested in the study.  |
| Tissue Type           | The type of tissue tested in the study.  |
| Timepoint             | The time period when the sample was taken.   |
| Select a Gene/Pathway | The gene, gene signature, or pathway of interest.<br>If you would like to run the analysis on the entire chip, leave this field blank.   |
| Select SNP Type       | Select the type of SNP data being used: <ul style="list-style-type: none"> <li>Genotype – Use for categorical variables.</li> <li>Copy Number – Use for continuous variables.</li> </ul> <p><b>Note:</b> This option is only available when you drag SNP data into the variable selection box.</p> <p><b>Note:</b> Both Genotype and Copy Number data may not be available for all studies involving SNP data.</p>   |
| Aggregate Probes?     | The checkbox can be selected if the variable chosen is either gene expression data or SNP copy number data.<br><br>If the checkbox is selected, the algorithm WGCNA (weighted correlation network analysis) is employed. For genes that are comprised of multiple probes, WGCNA selects the probe that best represents the overall expression level or copy number.<br><br><b>Note:</b> WGCNA was developed by the Department of Human Genetics at UCLA. For more information, see <a href="http://www.genetics.ucla.edu/labs/horvath/CoexpressionNetwork/">http://www.genetics.ucla.edu/labs/horvath/CoexpressionNetwork/</a> . |

9. Click **Apply Selections**.

10. Define any additional required variables, then click **Run**.

## Other Features

The sections below illustrate additional features in the Advanced Workflow tab.

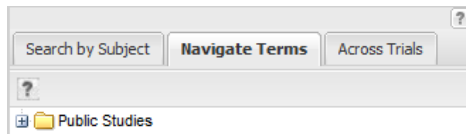
### Save to PDF

The Save to PDF feature allows you to save analyses run through the Advanced Workflow function within Dataset Explorer.

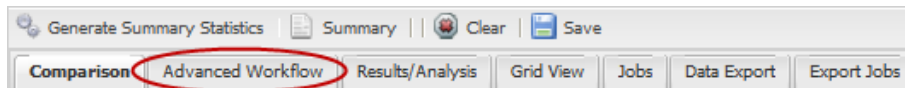
#### To save advanced workflow analyses as a PDF file:

1. Click the tranSMART **Dataset Explorer** tab to display the Dataset Explorer window.
2. In the left pane of the Dataset Explorer window, click the **Navigate Terms** tab.

The navigation tree appears, showing the categories of available studies:



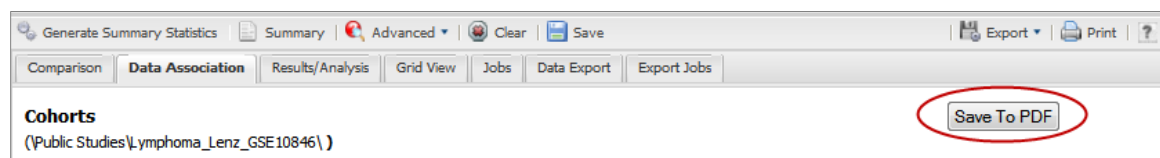
3. Select the study of interest.
4. Drag the study of interest into a subset definition box in Subset 1.
5. Click the **Advanced Workflow** tab above Subset 1:



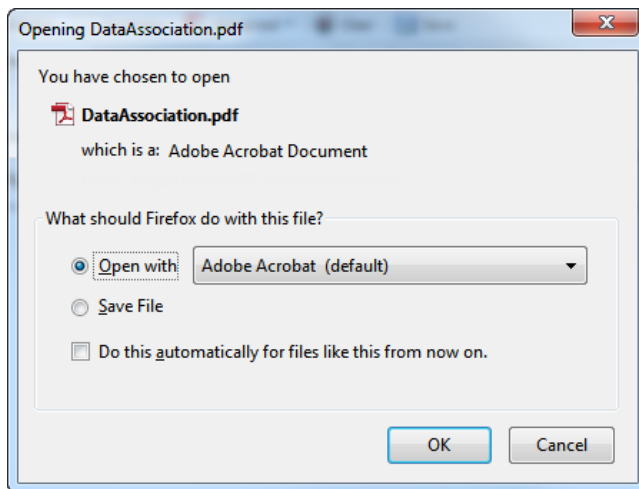
6. Select the analysis you wish to perform, and define the variables accordingly.
7. Click **Run**.

Your analysis appears below the variable selection panes.

8. Click **Save to PDF**:



The following dialog appears:



9. Select **Open with** or **Save File**, then click **OK**.

## Download Raw R Data

Analyses run through the Advanced Workflow tool within Dataset Explorer use R for computation. You are able to download raw R data for use in an external tool.

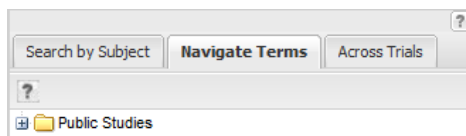


For more information on The R Project for Statistical Computing, visit the following site: [www.r-project.org](http://www.r-project.org).

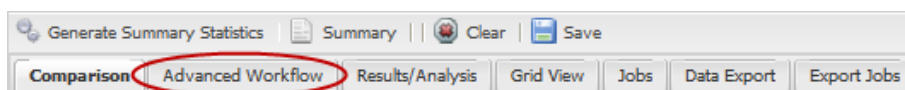
### To download advanced workflow analyses as raw R data:

1. Click the tranSMART **Dataset Explorer** tab to display the Dataset Explorer window.
2. In the left pane of the Dataset Explorer window, click the **Navigate Terms** tab.

The navigation tree appears, showing the categories of available studies:



3. Select the study of interest.
4. Drag the study of interest into a subset definition box in Subset 1.
5. Click the **Advanced Workflow** tab above Subset 1:



6. Select the analysis you wish to perform, and define the variables accordingly.

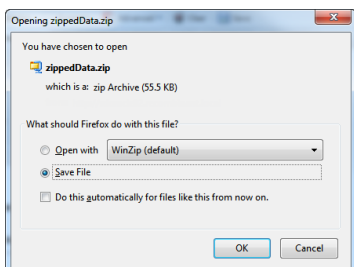
7. Click **Run**.

Your analysis appears below the variable selection panes.

8. Click **Download raw R data**:



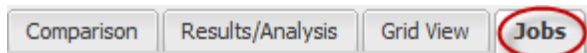
The following dialog appears:



9. Select whether you would like to open the file or save it to your hard drive, then click **OK**.

## The Jobs Tab

The Jobs tab allows you to review analyses you have run previously.



Each advanced workflow analysis that you have run in the past seven days is logged in the Jobs tab in a spreadsheet format.

The columns of information in the Jobs tab are described below:

| Column | Description  |
|--------|--|
| Name   | <p>The name of the analysis run. The format of the name is as follows:</p> <div style="display: flex; justify-content: center; align-items: center; gap: 10px;"> <div style="border: 1px solid black; padding: 2px; font-weight: bold; color: red;">tranSMART<br/>user name</div> <div style="border: 1px solid black; padding: 2px; font-weight: bold; color: red;">Type of<br/>analysis run</div> <div style="border: 1px solid black; padding: 2px; font-weight: bold; color: red;">Job ID</div> </div> <p style="text-align: center; color: blue; font-weight: bold; margin-top: 10px;"><u>user-PCA-5174</u></p> |

| Column     | Description   |
|------------|---|
| Status     | <p>The status of the analysis. Statuses are explained below:</p> <ul style="list-style-type: none"> <li>▪ <b>Completed</b> – The job has finished and a visualization or analysis is available.</li> <li>▪ <b>Started</b> – The job has been started and is still processing.</li> <li>▪ <b>Uploading File</b> – You have selected to load additional data into your visualization, and the data is still in the process of uploading to tranSMART.</li> <li>▪ <b>Error</b> – The job did not complete due to an error.</li> <li>▪ <b>Cancelled</b> – The job was cancelled and will not complete.</li> </ul> |
| Run Time   | The time the analysis took to process.  |
| Started On | The date and time that the analysis was started.  |



Click the **Refresh** button to view any changes that have been made since the Jobs tab initially populated.

## Viewing a Logged Job

Each advanced analysis that you have run in the previous seven days will be logged in the **Jobs** tab. You may view the visualization or analysis again by selecting it from the list.

### To run a logged advanced workflow:

1. Run tranSMART, then click the **Dataset Explorer** tab.
2. In the right pane, click the **Jobs** tab:



3. Click the hyperlink of the analysis you are interested in viewing:

| Name                              | Status         | Run Time      | Started On              |
|-----------------------------------|----------------|---------------|-------------------------|
| <a href="#">user-Compare-5221</a> | Completed      | 8.707 seconds | 2011-01-01 00:00:00.000 |
| <a href="#">user-Select-5207</a>  | Started        |               | 2011-01-01 00:00:00.000 |
| <a href="#">user-PCA-5179</a>     | Uploading file |               | 2011-01-01 00:00:00.000 |
| <a href="#">user-Select-5207</a>  | Error          |               | 2011-01-01 00:00:00.000 |
| <a href="#">user-PCA-5174 68</a>  | Started        |               | 2011-01-01 00:00:00.000 |
| <a href="#">user-PCA-5179</a>     | Error          |               | 2011-01-01 00:00:00.000 |

Refresh







## Chapter 4

# Sample Explorer

Sample Explorer lets you search for tissue and blood samples of interest so that you can learn more about the samples; for example, you can look up sample IDs and locate the study that produced the samples in the Dataset Explorer.

The Sample Explorer window has two panes:

- **Right pane – Select a primary search filter**

Lets you begin to search for samples. For information, see [Select a Primary Search Filter](#) below.

- **Left pane – Recent Updates**

Lists up to ten of the most recent sample updates in the database.

For information about a sample update, including the number and source of updated records, click the item in the list.

## Select a Primary Search Filter

---

This pane of the Sample Explorer window lets you initiate a search for samples by selecting the primary search filter. After you select a search filter, a second Sample Explorer window appears where you can view the search results and refine the search by selecting additional filters.

Search filters are organized in the following categories:

- **Data Type** – The type of data associated with the samples
- **Pathology** – The type of disease associated with the samples
- **Tissue** – The physical source of the samples, such as liver or colon tissue
- **Dataset** – The study that generated the samples

Note that the number of samples that are associated with a filter appear in parentheses after the filter name.

You can select a select a primary filter by searching or by browsing for the filter.

**To search for a primary search filter:**

1. Click the search filter category to search within, or accept the default of **All** categories:

Search for filter

Category: **ALL** | datatype | pathology | tissue | dataset | biobank

Search field

Search filter categories

2. Type part or all of the filter name into the **Search** field.

The search engine displays a dropdown list containing all the filters within the selected category that begin with the text you typed. For example, if you type the letter **G** in the **Search** field for an all-category search, you might see this:

Search for filter

Category: **ALL** | datatype | pathology | tissue | dataset | biobank

G

- Pathology> Gene Expression (1)
- Pathology> Gastric Cancer (186)
- Tissue> Gastric Mucosa (186)
- DataSet> GSE9285 (43)

Up to 20 filters can be listed. If the filter you want does not appear, type a more complete name in the **Search** field.

3. When the filter you want appears in the list, click the filter name.

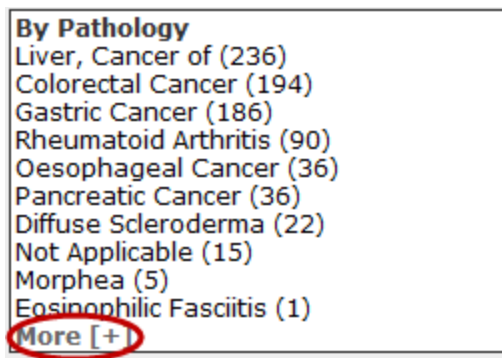
The search begins immediately, and the results are displayed in a new window (see [View and Refine Sample Search Results](#) on page 90).



You can only initiate a search by clicking a filter name in the dropdown list. You cannot initiate the search by typing the filter name and pressing the **Enter** key.

**To browse for a primary search filter:**

1. Click a filter name in one of the category browser boxes displayed below the search filter.
2. If you do not see the filter you want in a particular category, click **More** at the bottom of the box:

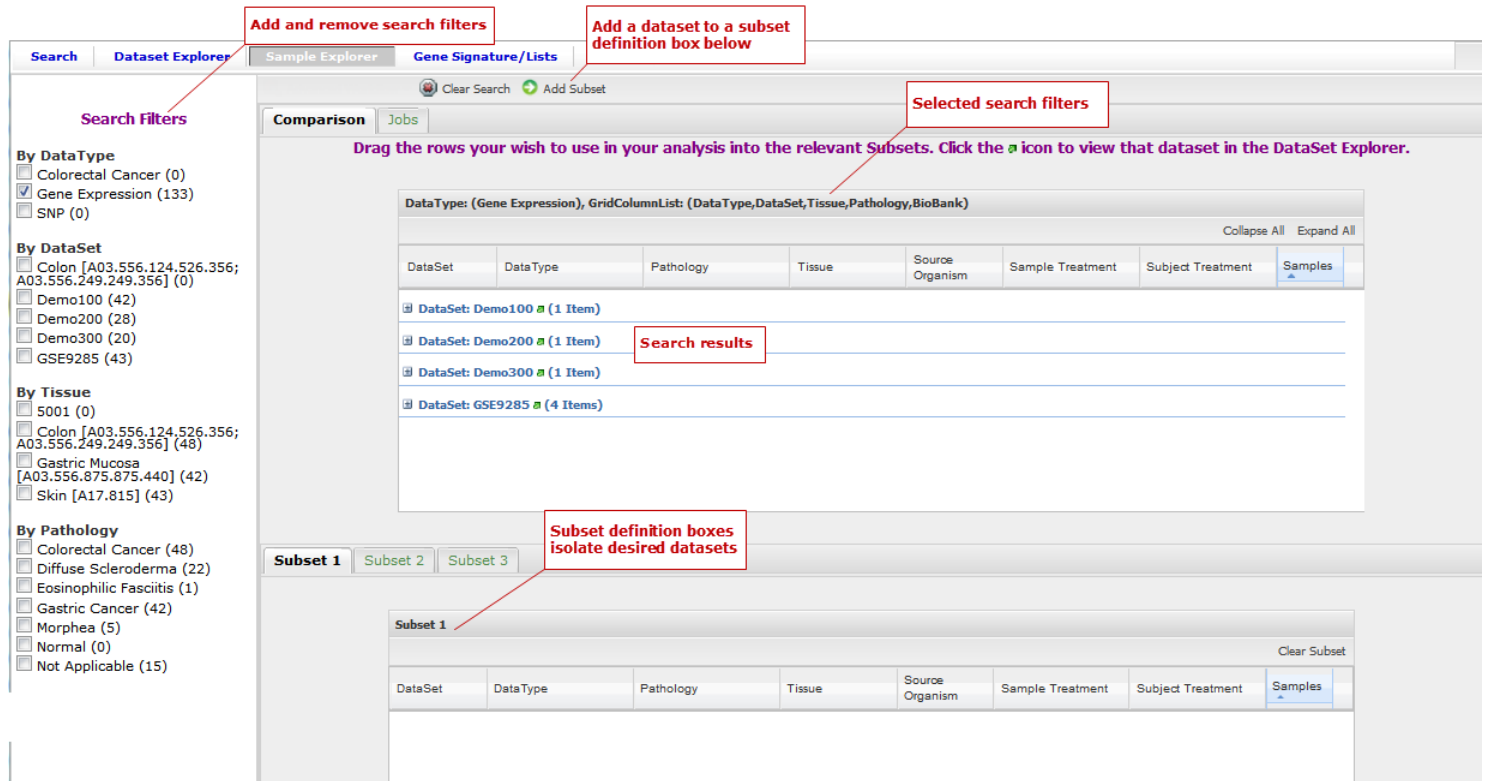


When you click a filter, the search begins immediately, and the results are displayed in a new window (see [View and Refine Sample Search Results](#) on page 90).

## View and Refine Sample Search Results

After you have selected a [primary search filter](#), a new Sample Explorer window appears, displaying the results of the search. The left pane of the window contains all the search filters, allowing you to narrow the search results.

The following figure illustrates the sections of this Sample Explorer window:



You can perform the following tasks in this Sample Explorer window:

- [Select and remove search filters](#)
- [Locate the study that produced the samples in the Dataset Explorer](#)
- [Project sample data onto a heatmap](#)
- [Re-sort the search results, and add/remove search result columns](#)

## Select and Remove Search Filters

You can refine a sample search result by adding and removing search filters, including the primary filter you initially selected. Search filters are listed in the left pane of the Sample Explorer window.

To select or remove a search filter, check or clear the check box next to the filter name.



Clicking a filter name rather than the check box next to the name will select that filter and deselect all currently selected filters.

The filters you select are joined together in a search string by the logical operators **AND** and **OR**, as follows:

- Filters within a filter category (such as **DataType**) are joined by **OR**.
- Filters in different filter categories are joined by **AND**.

For example, the search string for the filter selections illustrated below is:

```
(RBM OR Gene Expression) AND (Colorectal Cancer OR Gastric Cancer)
```

### By DataType

- ☒ RBM (90)
- ☒ Gene Expression (691)
- ☐ SNP (40)

### By Pathology

- ☐ Liver, Cancer of (236)
- ☒ Colorectal Cancer (194)
- ☒ Gastric Cancer (186)
- ☐ Rheumatoid Arthritis (90)
- ☐ Oesophageal Cancer (36)

## Locate the Source of the Samples in Dataset Explorer

If a dataset of samples was collected for a Dataset Explorer study, you can link back to the study to view information such as the study owner, study description and purpose, demographics of the participants, and other data relevant to the samples.



When you link back to a Dataset Explorer study, and then return to Sample Explorer, the filters you had previously selected in Sample Explorer are cleared.

### To link back to the associated Dataset Explorer study:

1. If the dataset of interest is not included in the result set, refine the search by selecting additional search filters (see [Select and Remove Search Filters](#) on page 91).
2. When the dataset of interest appears, click the dataset name in the **DataSet** column of the result set:

| DataSet                   | DataType        | Pathology      | Tissue                              |
|---------------------------|-----------------|----------------|-------------------------------------|
| DataSet: Demo100 (1 Item) |                 |                |                                     |
| <a href="#">Demo100</a>   | Gene Expression | Gastric Cancer | Gastric Mucosa<br>[A03.556.875.8... |

When you click a dataset link, the following actions occur automatically:

- a. Dataset Explorer opens.
- b. The dataset name you clicked is inserted into the **Search** field of the Dataset Explorer **Search By Subject** tab.
- c. The search is immediately executed, and one or more matching studies, or sub-nodes of studies, is listed below the **Search** field:

Search by Subject | Navigate Terms | Across Trials

Search:  AND Type:

Found 1 results.

Milano\_Scleroderma\_GSE9285

3. Open and explore the study of interest.

For information, see [Branches and Leaves of the Navigation Tree](#) on page 34.



If the study name is grayed out, or an Access Is Restricted warning is displayed when you try to open the study, you have not been granted access to the study. Contact a tranSMART administrator if you want to request access.



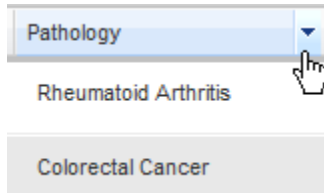
## Manage the Sample Search Result List

You can make the following adjustments to the search result list:

### Sort by Column

**To sort the result list by the contents of a column:**

1. Click the right side of the column heading to pull down the menu:

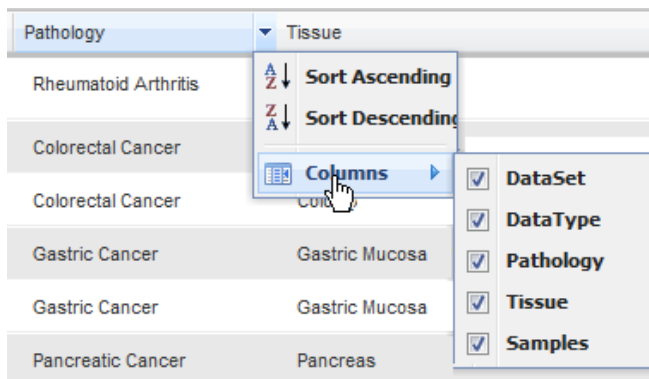


2. Click **Sort Ascending** or **Sort Descending**.

### Add and Remove Columns

**To add and remove columns:**

1. Click the right side of the column header to pull down the menu.
2. Hover the mouse pointer over Columns to display the submenu of column headings:



3. Check or clear the check boxes to add or remove a column from the search result.



If there are more rows in the result set than can be displayed at one time, a vertical scroll bar appears at the right of the result set. However, this scroll bar may be hidden from view. To check, move the horizontal scroll bar at the bottom of the window all the way to the right to expose the result set's vertical scroll bar. If the vertical scroll bar is not there, all the rows in the result set currently are displayed.



# Gene Signatures and Gene Lists

The tranSMART gene signature wizard guides you through the process of creating a gene signature or gene list. You specify whether the gene signature or list is publicly available to other tranSMART users or is reserved for your private use.

Once you create the gene signature or list, it can be used in tranSMART searches to find clinical studies and experiments where the differentially regulated genes overlap with the genes contained in the gene signature or list. This will generate a set of hypotheses about diseases or treatments that may have similar genes deregulated, and that can help you develop a further set of experiments.



This chapter uses the term “gene signature” to refer to both gene signatures and gene lists.

## Creating a Gene Signature

---

There are two basic tasks involved in creating a gene signature:

1. Add the list of genes for the gene signature to a text file.  
Genes can be indicated by gene symbol or by their associated probe set ID.
2. Use the gene signature wizard to define the information on which the gene signature is based, such as species, source of data, and test type, and also to import into the gene signature definition the text file containing the genes.

### Step 1. Adding the Genes to a Text File

The gene signature wizard expects to import the genes for the gene signature from a tab-separated text file. The file must contain one, and possibly two, columns of information:

- First column – A list of gene symbols or probe set IDs.
- Optional second column – The fold change ratios associated with the gene symbols or probe set IDs.

The fold change ratios can be either **actual values** (for example, 12.8 or -12.8) or one of the following **composite values**:

- ☐ **-1**. All down-regulated gene expressions.
- ☐ **1**. All up-regulated gene expressions.
- ☐ **0**. No change.

The following table shows the different ways you can specify the genes for your gene signature:

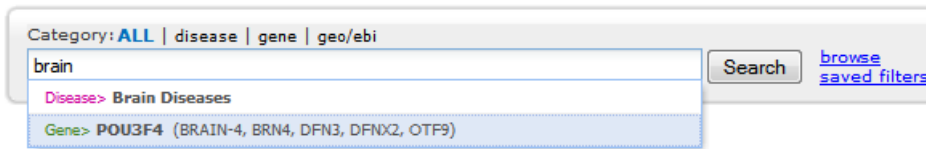
| Contents of File                        | Format                                     | Examples  |
|---|--|---|
| Gene symbols only                       | <i>GeneSymbol</i>                          | TCN1<br>IL1RN<br>KIAA1199<br>G0S2   |
| Gene symbols,<br>actual fold change     | <i>GeneSymbol</i> <tab> <i>ActualFC</i>    | CXCL5    -19.19385797<br>IL8RB    -18.21493625<br>FPR1     -17.6056338<br>FCGR3A   -15.69858713   |
| Gene symbols,<br>composite fold change  | <i>GeneSymbol</i> <tab> <i>CompositeFC</i> | CXCL5    -1<br>IL8RB    -1<br>MMP3     0<br>SOD2     1  |
| Probe set IDs only                      | <i>ProbesetID</i>                          | 224301_x_at<br>1398191_at<br>Dr.2473.1.A1_at<br>A_24_P93251   |
| Probe set IDs,<br>actual fold change    | <i>ProbesetID</i> <tab> <i>ActualFC</i>    | 224301_x_at    -<br>19.19385797<br>1398191_at    -<br>18.21493625<br>Dr.2473.1.A1_at -<br>17.6056338<br>A_24_P93251    -<br>15.69858713 |
| Probe set IDs,<br>composite fold change | <i>ProbesetID</i> <tab> <i>CompositeFC</i> | 224301_x_at    -1<br>1398191_at     0<br>Dr.2473.1.A1_at 1<br>A_24_P93251    -1   |

## Using tranSMART to Select Genes

You can use the tranSMART Search tool to help you select the list of genes for your gene signature. For example, suppose you are interested in brain diseases, and want to create a gene signature consisting of genes that were strongly up-regulated in an experiment involving Alzheimer's patients.

You can use tranSMART to select the genes for the gene signature as follows:

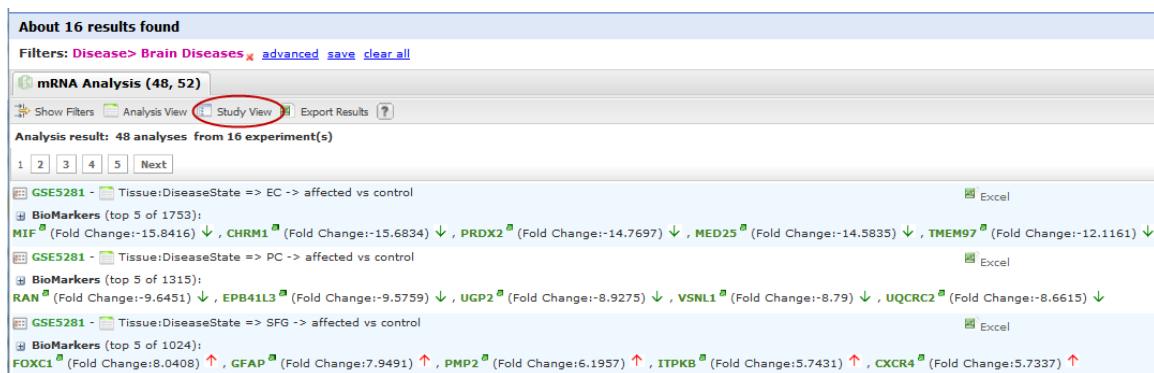
1. In tranSMART, click the **Search** tab to display the Search window.
2. Type **brain** in the Search field:




3. Click **Disease> Brain Diseases**.

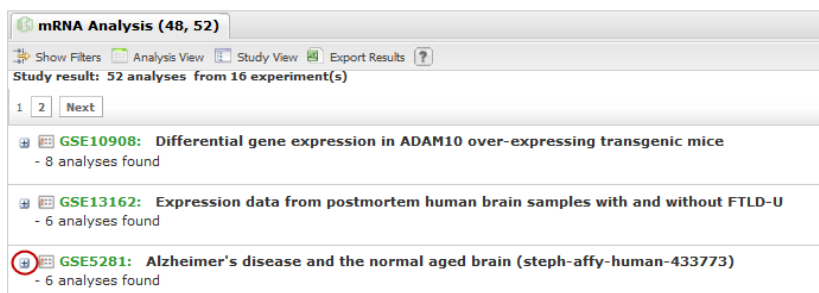
In a few seconds, the search result appears.

4. Click the **mRNA Analysis** tab, then click the **Study View** button:



tranSMART displays a list of all the experiments related to brain diseases.

5. Scroll through the list of experiments until you find the one to use as the basis of your gene signature.
6. Click the **+** icon (  ) to the left of the experiment name:



A list of the analyses based on this experiment appears.

- Click the **Excel** button for the analysis that you want to use for your gene signature:

**GSE5281: Alzheimer's disease and the normal aged brain (steph-affy-human-433773)**  
- 6 analyses found

Tissue:DiseaseState => MTG -> affected vs control Excel

**BioMarkers** (top 5 of 14067):  
**LIFR** (Fold Change:21.7284) ↑, **MALAT1** (Fold Change:18.8822) ↑, **MAFF** (Fold Change:17.8537) ↑, **VSNL1** (Fold Change:-16.5996) ↓, **IGFBP7** (Fold Change:16.0599) ↑

Tissue:DiseaseState => PC -> affected vs control Excel

**BioMarkers** (top 5 of 13486):  
**RAN** (Fold Change:-9.6451) ↓, **EPB41L3** (Fold Change:-9.5759) ↓, **UGP2** (Fold Change:-8.9275) ↓, **VSNL1** (Fold Change:-8.79) ↓, **UQCRC2** (Fold Change:-8.6615) ↓

Tissue:DiseaseState => HIP -> affected vs control Excel

**BioMarkers** (top 5 of 11448):  
**YWHAH** (Fold Change:-11.455) ↓, **EP400** (Fold Change:9.7391) ↑, **ARPC1A** (Fold Change:-9.0962) ↓, **HSPA8** (Fold Change:-8.1439) ↓, **SCAMP1** (Fold Change:7.8051) ↑

Tissue:DiseaseState => EC -> affected vs control Excel

**BioMarkers** (top 5 of 11178):  
**MIF** (Fold Change:-15.6834) ↓, **CHRM1** (Fold Change:-14.7697) ↓, **MED25** (Fold Change:-14.5835) ↓, **TMEM97** (Fold Change:-12.1161) ↓

Tissue:DiseaseState => VCX -> affected vs control Excel

**BioMarkers** (top 5 of 11178):  
**VSNL1** (Fold Change:-5.5248) ↓, **PENK** (Fold Change:-4.5258) ↓, **NAP1L5** (Fold Change:-4.4647) ↓, **TAC1** (Fold Change:-4.2787) ↓, **TBC1D5** (Fold Change:-4.2102) ↓

Tissue:DiseaseState => SFG -> affected vs control Excel

**BioMarkers** (top 5 of 9455):  
**FOXCl** (Fold Change:8.0408) ↑, **GFAP** (Fold Change:7.9491) ↑, **PMP2** (Fold Change:6.1957) ↑, **ITPKB** (Fold Change:5.7431) ↑, **CXCR4** (Fold Change:5.7337) ↑

This action exports the analysis information, including the gene expression data, to a Microsoft Excel file.

- Click **Open** in the File Download dialog.

Excel starts up and displays the analysis data – for example:

|   | A                             | B           | C                 | D       | E                | F           | G      |
|---|-------------------------------|-------------|-------------------|---------|------------------|-------------|--------|
| 1 | Analysis                      | ProbeSet    | Fold Change Ratio | p-Value | adjusted p-value | TEA p-Value | Gene   |
| 2 | Tissue:DiseaseState => SFG -> | 206100_at   | 1.3846            | 0.0006  | 0.0048           | 0.13        | CPM    |
| 3 | Tissue:DiseaseState => SFG -> | 206165_s_at | -1.1092           | 0.0303  | 0.095            | 0.1816      | CLCA2  |
| 4 | Tissue:DiseaseState => SFG -> | 206171_at   | 1.3041            | 0.0038  | 0.0206           | 0.1439      | ADORA3 |
| 5 | Tissue:DiseaseState => SFG -> | 206268_at   | -1.3401           | 0.0073  | 0.0339           | 0.1376      | LEFTY1 |
| 6 | Tissue:DiseaseState => SFG -> | 206452_x_at | 1.2725            | 0.0079  | 0.0358           | 0.1497      | PPP2R4 |
| 7 | Tissue:DiseaseState => SFG -> | 206597_at   | -1.147            | 0.0753  | 0.1821           | 0.1739      | NRL    |

- Remove any rows containing genes that you do not want to include in the gene signature.
- Remove the column headings and all data from the spreadsheet except for the gene symbol and fold change ratio, then export the remaining data to a tab-separated text file using the Excel **Save as type** option **Text (Tab delimited) (\*.txt)**.

## Step 2. Creating the Gene Signature

1. In tranSMART, click the **Gene Signature/Lists** tab.
2. Click the **New Signature** button.

The first page of the gene signature wizard appears:

**Gene Signature Create**



Instructions ▼

**Page 1: Definition:**

Signature/List Name\*

Description

Note, the creator of this signature will be 'Anthony Ioven' at the current system time

 Meta-Data  Cancel



Required fields on gene signature wizard pages are marked with a red asterisk (\*).

You can find additional information about the gene signature wizard by clicking **Information** on any wizard page.

3. Specify a name (required) and an optional description for your gene signature, then click Meta-Data to proceed to the next gene wizard page.

The second page of the gene signature wizard appears:

**Gene Signature Create**

**Instructions ▼**

**Page 2: Meta-Data:**

|                         |   |                      |
|-------------------------|---|----------------------|
| Source of list          | select source ▼                                       |                      |
| Owner of data           | select owner of the data ▼                            |                      |
| Stimulus                | i.e. LPS, polyIC, etc:                                | <input type="text"/> |
|                         | Dose, units, and time:                                | <input type="text"/> |
| Treatment               | Drug treatment used in assay:                         | <input type="text"/> |
|                         | Dose, units, and time:                                | <input type="text"/> |
|                         | <b>OR Enter:</b>                                      |                      |
|                         | J&J Compound:   | select compound ▼    |
|                         | Protocol Number:                                      | <input type="text"/> |
| PMIDs (comma separated) | <input type="text"/>                                  |                      |
| Species*                | select relevant species ▼                             |                      |
| Technology Platform*    | select tech platform ▼                                |                      |
| Tissue Type             | select relevant tissue ▼                              |                      |
| Experiment Type         | select experiment type ▼                              |                      |
|                         | If applicable, ATCC designation: <input type="text"/> |                      |

Definition Next Cancel



4. Specify values in the required fields **Species** and **Technology Platform**, and also in any other relevant fields, then click Next to proceed to the final gene signature wizard page:

**Gene Signature Create**

Instructions ▼

**Page 3: Analysis Meta-Data:**

Analysis Performed By:

Normalization Method:

Analysis Info

Category:

Method:

Multiple Testing Correction Employed? ☐ Yes ☐ No

P-value Cutoff\*:

**File Upload Information (tab delimited text only, no .xls Excel files):** [See Samples](#)

File Information\*

File schema:

Fold change metric:

Upload File\* (tab delimited text files only)

5. Specify values in the required field **P-value Cutoff**.
6. In the section **File Upload Information**, describe the text file you created in the section [Step 1. Adding the Genes to a Text File](#) on page 95, using the required fields **File Information** and **Upload File**:
- In the **File schema** section of **File Information**, select **Gene Symbol <tab> Metric Indicator** or **Probe Set Symbol <tab> Metric Indicator**, depending on the method you chose to specify the genes.
  - In the **Fold change metric** section of **File Information**, select one of the following choices from the dropdown:


| Fold Change Metric Indicator                  | Description  |
|---|--|
| Actual fold change                            | The text file contains actual fold change values for each gene symbol or probe set ID.   |
| Not used                                      | The text file contains gene symbols or probe set ID only. There are no associated fold change values.  |
| -1 (down), 1 (up), 0 (optional for unchanged) | The fold change values are not actual values. They simply represent whether the gene expression was down-regulated (-1), up-regulated (1), or unchanged (0). |

- In **Upload File**, specify the path and name of the file that contains the genes to import. Use the **Browse** button to select the file from the navigation tree.

- Specify values in any other relevant fields on this gene wizard page, then click **Save** to save the gene signature.

The new gene signature appears in the **Gene Signature List** at the top of the Gene Signature/List view:

#### Gene Signature List

| My Signatures (1) ▲   |                  |              |         |               |             |        |           |         |                |                  |
|---|------------------|--------------|---------|---------------|-------------|--------|-----------|---------|----------------|------------------|
| Name  | Author           | Date Created | Species | Tech Platform | Tissue Type | Public | Gene List | # Genes | # Up-Regulated | # Down-Regulated |
|  Trainee9 Gene Signature | Training Account | 2009-08-08   | Human   | GPL8300       | Lung        | No     | No        | 18      | 7              | 11               |

-- Select Action -- ▼


## Making a New Gene Signature Public

By default, a newly created gene signature is private.

### To make a gene signature public:

- In the **Gene Signature List**, click the **Select Action** dropdown to the right of the gene signature you just created.
- Click **Make Public** in the dropdown list:


#### Gene Signature List

| My Signatures (1) ▲   |                  |              |         |               |             |        |           |         |                |                  |
|---|------------------|--------------|---------|---------------|-------------|--------|-----------|---------|----------------|------------------|
| Name  | Author           | Date Created | Species | Tech Platform | Tissue Type | Public | Gene List | # Genes | # Up-Regulated | # Down-Regulated |
|  Trainee9 Gene Signature | Training Account | 2009-08-08   | Human   | GPL8300       | Lung        | No     | No        | 18      | 7              | 11               |

-- Select Action -- ▼  
 -- Select Action --  
 Clone  
 Delete  
 Edit  
 Edit Items  
 Excel Download  
 Make Public

After you click **Make Public**, the value in the **Public** column for the gene signature changes from **No** to **Yes**:

#### Gene Signature List

| My Signatures (1) ▲   |                  |              |         |               |             |        |           |         |                |                  |
|---|------------------|--------------|---------|---------------|-------------|--------|-----------|---------|----------------|------------------|
| Name  | Author           | Date Created | Species | Tech Platform | Tissue Type | Public | Gene List | # Genes | # Up-Regulated | # Down-Regulated |
|  Trainee9 Gene Signature | Training Account | 2009-08-08   | Human   | GPL8300       | Lung        | Yes    | No        | 18      | 7              | 11               |



transSMART users assigned the role `ROLE_ADMIN` have access to both public and private gene signatures.


## Performing Actions on Your Gene Signatures

To edit or perform other actions on a gene signature in your gene signature list:


1. In tranSMART, click the **Gene Signature/Lists** tab.

The **Gene Signature List** appears, containing all the genes you have created:

### Gene Signature List

| My Signatures (1) ▲   |                  |              |         |               |             |        |           |         |                |                       |
|---|------------------|--------------|---------|---------------|-------------|--------|-----------|---------|----------------|-----------------------|
| Name  | Author           | Date Created | Species | Tech Platform | Tissue Type | Public | Gene List | # Genes | # Up-Regulated | # Down-Regulated      |
|  Trainee9 Gene Signature | Training Account | 2009-08-08   | Human   | GPL8300       | Lung        | Yes    | No        | 18      | 7              | 11                    |
|   |                  |              |         |               |             |        |           |         |                | -- Select Action -- ▼ |

2. Click the **Select Action** dropdown for the gene signature you are acting on. The dropdown contains all the actions you can perform on the gene signature:

| Action         | Description  |
|----------------|--|
| Clone          | <p>Create an exact duplicate of the gene signature definition (<i>except</i> for the text file containing the gene symbols and fold change values), and display the definition in the gene signature wizard.</p> <p>Cloning a gene signature helps you create a new gene signature with a similar definition to an existing one. However, it is expected you will import a different set of genes into the gene signature.</p>   |
| Delete         | Delete the gene signature.   |
| Edit           | <p>Open the gene signature in the gene signature wizard for editing.</p> <p>The gene signature wizard displays all the information in the gene signature, including the reference to the text file containing the list of genes and fold change values. If you want to choose a different text file, click the following label:</p> <p></p> <p>To save any changes you make during editing, you must click the <b>Save</b> button on the third page of the wizard.</p> |
| Edit Items     | Add, delete, or modify one or more genes in the text file containing the gene symbols and fold change values.  |
| Excel Download | <p>Generate the entire contents of the gene signature, including the information in the text file containing the gene symbols and fold change values, to a Microsoft Excel spreadsheet.</p> <p>The gene signature definition and gene symbols/fold change values are written to separate spreadsheets.</p>   |

| Action      | Description   |
|-------------|---|
| Make Public | <p>Make a private gene signature public.</p> <p><b>Note:</b> To make a public gene signature private, edit the gene signature and set the <b>Public?</b> field to <b>No</b> on the first page of the gene signature wizard:</p> <p><b>Public?</b> <input type="radio"/> Yes <input checked="" type="radio"/> No</p> |

## Performing Actions on Other Users' Signatures

You can perform actions on gene signatures that other tranSMART users have created. The gene signatures you can access and the actions you can perform on them depend on the role assigned to your tranSMART user ID, as follows:

| Role  | Authorized Actions  |
|---|---|
| ROLE_ADMIN  | All actions on all gene signatures, both public and private.                      |
| ROLE_SPECTATOR<br>ROLE_STUDY_OWNER<br>ROLE_DATASET_EXPLORER_ADMIN | Only <b>Clone</b> and <b>Excel Download</b> , and only on public gene signatures. |

### To edit or perform actions on a gene signature other than your own:

1. In tranSMART, click the **Gene Signature/Lists** tab.
2. Click **Public Signatures** to open the list of public gene signatures:

#### Gene Signature List

| My Signatures (1) ▲         |                  |              |         |               |             |        |           |         |                |                  |
|-----------------------------|------------------|--------------|---------|---------------|-------------|--------|-----------|---------|----------------|------------------|
| Name                        | Author           | Date Created | Species | Tech Platform | Tissue Type | Public | Gene List | # Genes | # Up-Regulated | # Down-Regulated |
| Traineer9<br>Gene Signature | Training Account | 2009-08-08   | Human   | GPL8300       | Lung        | No     | No        | 18      | 7              | 11               |
| -- Select Action -- ▼       |                  |              |         |               |             |        |           |         |                |                  |

Public Signatures (11) ▼




tranSMART users assigned the role `ROLE_ADMIN` will see **Other Signatures** instead of **Public Signatures**.

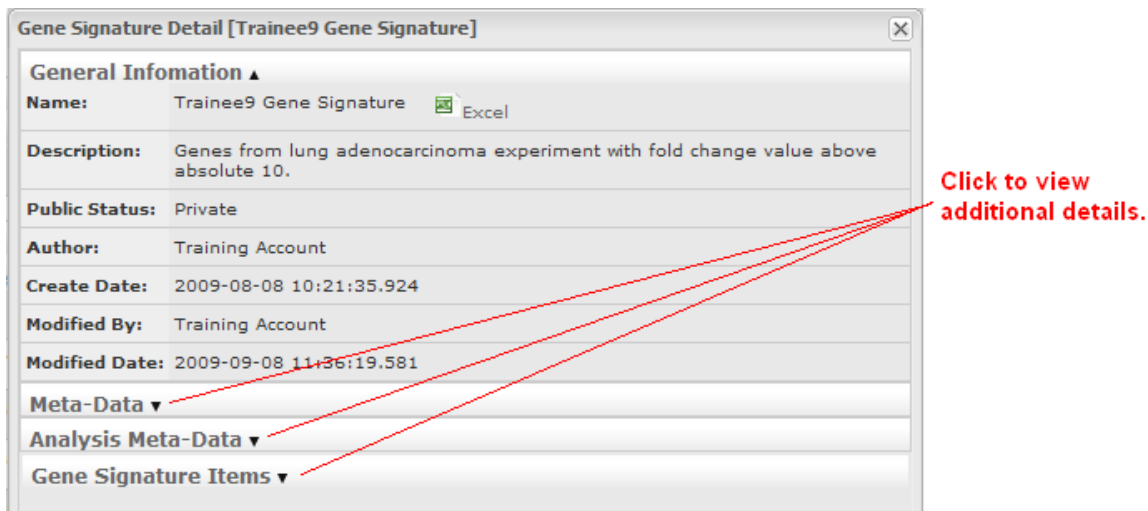
3. Click the **Select Action** dropdown for the gene signature you want to act on.
4. Select the action you want to perform on the gene signature.

## Viewing a Gene Signature Definition

You can view the definition of a gene signature, including its list of genes and fold change values, for any gene signature you are authorized to access.

To view a gene signature definition, click the **Detail** icon (  ) next to the gene signature name.

The Gene Signature Detail dialog appears, containing the gene signature definition:





## Chapter 6

# Other Tasks

In addition to Search, Dataset Explorer, and Gene Signature/List, the tranSMART toolbar includes the following tool tabs:

- **Request Consult**

Email a request for information that cannot be found in the data warehouse.

- **Feedback**

Email questions, problem reports, enhancement requests, or any other feedback about the tranSMART application.

- **Help**

Display links to the tranSMART online and PDF documentation sets.

- **Admin**

Perform user administration functions such as creating users, assigning roles to users, and enabling or disabling user accounts.

This tool tab is visible only to users who are assigned the role `ROLE_ADMIN`.





## Appendix A

# How TEA Scores Are Calculated

This appendix summarizes the operations transSMART performs to calculate the overall TEA score for an experiment, and the data inputs that the calculation requires. Pseudocode representations of the operations being performed are included where they may clarify the operation.

## Data Inputs to the TEA algorithm

---

One of the following:

- A gene signature or gene list containing any number of genes, with a binary up- or down-regulation flag based on fold change.
- A pathway containing any number of genes.

And:

- A gene search result list for each signature, list, or pathway gene. Result lists contain experimental comparisons.

## Operations

---

1. Compute the average fold change and standard deviation for all genes in the comparison.
2. Compute a normalized p-value (NPV) for each gene in the comparison, based on its fold change (fc) value, and the above average (ave) and standard deviation (std) values. Use a normal distribution function (CDF):

```
if (fc > 0)
  NPV = 1.0 - CDF(fc, ave, std)
else
  NPV = 1.0 - CDF(-fc, ave, std)
if NPV < 1.0e-15, set to 1.0e-15
```

3. For each gene in the gene signature, list, or pathway, search against experimental comparisons and extract those comparisons where the gene's normalized p-value is less than 0.05. This returns a comparison list.

4. Iterate through the comparison list. For each comparison (*c*), add the normalized p-value to one of two arrays of sums (*pv\_sum*), as follows:

- For gene signatures and gene lists, add the gene's normalized p-value to:
  - *pv\_sum*(*C*, up) if the gene's fold change in the signature (*sfc*) *and* in the comparison (*cfc*) are co-regulated.
  - *pv\_sum*(*C*, down) if the gene's fold change in the signature *and* in the comparison are anti-regulated.
- For pathways, add the gene's normalized p-value to:
  - *pv\_sum*(*C*, up) if the gene's comparison fold change (*cfc*) is up-regulated.
  - *pv\_sum*(*C*, down) if the gene's comparison fold change is down-regulated.

Also, use the logarithm of the normalized p-value to make the final TEA score more human readable:

```
if (gene_signature OR gene_list)
  if ( (sfc > 0 AND cfc > 0) OR (sfc < 0 AND cfc < 0) )
    pv_sum(C, up) += -Math.log(NPV)
    pv_count(C, up)++
  else
    pv_sum(C, down) += -Math.log(NPV)
    pv_count(C, down)++

if (gene_pathway)
  if (cfc > 0)
    pv_sum(C, up) += -Math.log(NPV)
    pv_count(C, up)++
  else
    pv_sum(C, down) += -Math.log(NPV)
    pv_count(C, down)++
```

5. Compute the min-LogP average (*pv\_ave*) for each sum:

```
pv_ave(C, up) = Math.exp(-pv_sum(C,up) / pv_count(C,up) )
pv_ave(C, down) = Math.exp(-pv_sum(C,down) / pv_count(C,down) )
```

6. Compute a TEA score (*pv\_tea*) for each min-LogP average through a binomial distribution function:

```
pv_tea(C, up) = 1.0 - Binom( N, pv_count(C,up), pv_ave(C,up) )
pv_tea(C, down) = 1.0 - Binom( N, pv_count(C,down), pv_ave(C,down) )
```

## Result

---

- **TEA score:** For gene signatures, lists, and pathways, the final TEA score is the more significant  $p_{v\_tea}$  value (the lower of the two  $p_{v\_tea}$  values).
- A gene signature or list is determined to be co-regulated or anti-regulated as follows:
  - **Co-regulated:** The more significant  $p_{v\_tea}$  value was derived from the sums associated with co-regulated fold change values ( $p_{v\_sum}(C, up)$ ).
  - **Anti-regulated:** The more significant  $p_{v\_tea}$  value was derived from the sums associated with anti-regulated fold change values ( $p_{v\_sum}(C, down)$ ).
- A pathway is determined to be up-regulated or down-regulated as follows:
  - **Up-regulated:** The more significant  $p_{v\_tea}$  value was derived from the sums associated with up-regulated fold change values ( $p_{v\_sum}(C, up)$ ).
  - **Down-regulated:** The more significant  $p_{v\_tea}$  value was derived from the sums associated with down-regulated fold change values ( $p_{v\_sum}(C, down)$ ).

Result

## Appendix B

# Rules for Loading OmicSoft Data

The following rules determine whether OmicSoft data is loaded into tranSMART databases:

### Rules for loading data into table **BIO\_ASSAY\_ANALYSIS\_DATA**

Load the data if one of the following two criteria is satisfied:

- fold\_change\_ratio is  $\geq 1.0$  OR  $\leq -1.0$   
AND  
preferred\_pvalue is null OR  $\leq 0.1$
- fold\_change\_ratio is null  
AND  
At least one of the following values is not null:
  - r\_value  
Pearson product-moment correlation coefficient.
  - rho\_value  
Spearman rank correlation coefficient
  - cut\_value
  - results\_value

### Rules for loading data into table **BIO\_ASSAY\_ANALYSIS\_DATA\_TEA**

Load the data if:

TEA\_NORMALIZED\_PVALUE  $\leq 0.05$

For information about the TEA algorithm, see [Appendix A: How TEA Scores Are Calculated](#).



## Appendix C

# Glossary of Terms

### AGGREGATE PROBES

Used in Dataset Explorer, the Aggregate Probes checkbox allows you to group probes used in high-dimensional data samples to form a total quantity that analyses will be performed on.

### ANALYSIS OF VARIANCE (ANOVA)

Analysis of Variance (ANOVA) is a statistical method used in Dataset Explorer to make concurrent comparisons between two or more means in a box plot.

### ANALYSIS VIEW

Used in the Search tool, the Analysis View option displays the statistically significant analyses from your search filter(s).

### ANTI-REGULATION

An analysis of a statistically significant experiment returned from a search against a gene signature or list is designated as *co-regulated* or *anti-regulated*.

### ARRAY DATA

See: [Microarray](#)

### ARRAYEXPRESS

Database of gene expression and other microarray data at the European Bioinformatics Institute (EBI).

See <http://www.ebi.ac.uk/arrayexpress> for details.

### BINOMIAL DISTRIBUTION

Graph that displays the discrete probability distribution of obtaining  $n$  successes out of  $N$  Bernoulli trials.

See <http://mathworld.wolfram.com/BinomialDistribution.html> for details.

## BIOMARKER

Short for Biological Marker, a biomarker is a key molecular or cellular event that links a specific environmental exposure to a health outcome.

## BOX PLOT

Also known as a Box and Whisker Plot, a box plot is a histogram-like method of displaying data. Box plots are useful when conveying location and variation information in datasets.

## CATEGORICAL VARIABLE

Also known as a nominal value, a categorical variable is one that has two or more categories, but with no intrinsic ordering to the categories. An example of a categorical value is hair color – there is no way to order these variables from highest to lowest.

## CENSORING VALUE

Used in Survival Analyses. The Censoring Value specifies which patients had the event whose time is being measured. For example, if the Time variable selected is Overall Survival Time (Years), an appropriate censoring variable is Patient Death.

## CHI SQUARED

Let the probabilities of various classes in a distribution be  $p_1, p_2, \dots, p_k$ , with observed frequencies  $m_1, m_2, \dots, m_k$ . The quantity

$$\chi^2 = \sum_{i=1}^k \frac{(m_i - N p_i)^2}{N p_i}$$

is therefore a measure of the deviation of a sample from expectation, where  $N$  is the sample size.

## COHORT

A group of subjects who have shared a specific event or characteristic.

## CONTINUOUS VARIABLE

Continuous variables have an infinite number of values between two points. For example, age or temperature.



**CO-REGULATION**

An analysis of a statistically significant experiment returned from a search against a gene signature or list is designated as *co-regulated* or *anti-regulated*.

**CORRELATION ANALYSIS**

A type of Regression Analysis, correlation analysis measures the correlation coefficient – the linear association between two variables. Values of the correlation coefficient are always between -1 and +1. A correlation coefficient of +1 indicates that two variables are perfectly related in a positive linear sense, while a correlation coefficient of -1 indicates that two variables are perfectly related in a negative linear sense.

**COX COEFFICIENT**

The Cox coefficient refers to the coefficients in a Cox regression model (also known as the proportional hazards model for survival-time). The analysis investigates the effects of one or more variables upon the time a specified event takes to happen. The cox coefficient relates to a hazard; a positive coefficient indicates a worse prognosis, while a negative coefficient indicates a protective effect of the variable.

**DATA BINNING**

Defers to a data pre-processing technique used to reduce observation errors and to allow continuous variables to become categorical. Clusters of data are replaced by a value representative of that cluster (often but not necessarily, the central value).

**DATA WAREHOUSE**

A database used for reporting and analysis.

**DATASET**

Collection of data, most commonly presented in a tabular form where each column represents a specific variable, and each row represents a value for that variable.

**DATASET EXPLORER**

Dataset Explorer lets you compare data generated for test subjects in two different study groups, based on criteria and points of comparison that you specify. Dataset Explorer is useful to help you test a hypothesis that involves the criteria and points of comparison that you select.

### **DEPENDENT VARIABLE**

In an experiment, the dependent variable is the response that is measured.

### **DIFFERENTIAL MODULATION**

### **DOWN-REGULATION**

An analysis of a statistically significant experiment returned from a search against a pathway is designated as *up-regulated* or *down-regulated*.

### **ENTREZ GENE**

Reference sequences for a wide range of species. For details, see <http://www.ncbi.nlm.nih.gov/gene/>.

### **ENTREZ GLOBAL**

Federated search engine that allows users to search various health sciences databases at the National Center for Biotechnology Information (NCBI) website.

See [www.ncbi.nlm.nih.gov/Entrez/](http://www.ncbi.nlm.nih.gov/Entrez/) for details.

### **FOLD CHANGE RATIO**

A number describing how much a quantity changes going from an initial to a final value. An initial value of 50 and a final value of 100 corresponds to a fold change of 2 (a two-fold increase).

### **GENE**

Stretches of DNA and RNA that code for a polypeptide or for an RNA chain – contains hereditary molecular information.

### **GENE CHIP**

See: [Microarray](#)

### **GENE EXPRESSION**

The flow of genetic information from gene to protein; the process, or the regulation of the process, by which the effects of a gene are manifested; the manifestation of a heritable trait in an individual carrying the gene or genes that determine it.

**GENE EXPRESSION OMNIBUS**

GEO is an international public repository that archives and freely distributes microarray, next-generation sequencing, and other forms of high-throughput functional genomics data submitted by the research community. For more information, see <http://www.ncbi.nlm.nih.gov/geo>.

**GENE SET ENRICHMENT ANALYSIS (GSEA)**

Computational method that determines whether an a priori defined set of genes shows statistically significant, concordant differences between two biological states (for example, phenotypes).

See <http://www.broadinstitute.org/gsea/index.jsp> for details.

**GENE SIGNATURE**

A group of genes whose combined expression pattern is uniquely characteristic of a medical condition or other clinical outcome of interest.

**GENE SYMBOL**

A unique abbreviation of a gene name consisting of italicized uppercase Latin letters and Arabic numbers. we use Entrez as the full list of genes (related to but not identical to HUGO)

See <http://www.genenames.org/> for details.

**GENECARDS**

Database that offers information about human genes (and mouse homologues).

See <http://www.genecards.org> for details.

**GOOGLE SCHOLAR**

Google application that provides a search of scholarly literature across multiple disciplines and sources.

See <http://scholar.google.com> for details.

**GPL PLATFORM**

A Platform record is composed of a summary description of the array or sequencer and, for array-based Platforms, a data table defining the array template. Each Platform record is assigned a unique and stable GEO accession

number (GPLxxx). A Platform may reference many Samples that have been submitted by multiple submitters.

### **HEATMAP**

Display of differential expression. Individual values contained in the matrix are represented by colors.

### **HIERARCHICAL CLUSTERING**

Hierarchical clustering is a type of clustering analysis whose goal is to organize data so that the objects in the same cluster are more similar to each other than to those in other clusters.

### **HIGH DIMENSIONAL DATA**

Datasets where the intersection of a subject and measurement is comprised of hundreds or thousands of points. For example, in a low dimensional data measurement such as height the intersection of subject and measurement is one number (ex. 180 cm) whereas in a high dimensional data measurement such as gene expression in a lymph node the measurement is 50,000 individual probe expression values.

### **HISTOGRAM**

A visual representation of the distribution of data values within a dataset.

### **HOMOLOGY**

The basis for comparative biology – where organs/structures from one organism are compared to a similar organ/structure in a different organism.

### **IN VITRO STUDY**

Those that are conducted using components of an organism that have been isolated from their usual biological surroundings.

### **IN VIVO STUDIES**

Experimentation using a whole, living organism.

### **INDEPENDENT VARIABLE**

In an experiment, the independent variable is the variable that is manipulated.

**JOB**

In tranSMART, a job refers to a command you have given Dataset Explorer to process or export data. Jobs and job-related events can be found within the **Jobs** tab in Dataset Explorer.

**KENDALL CORRELATION**

Kendall's rank correlation provides a distribution-free test of independence and a measure of the strength of dependence between two variables.

**K-MEANS CLUSTERING**

The K-Means clustering heatmap clusters genes and/or samples into a specified number of clusters. The result is  $k$  clusters, each centered around a randomly-selected data point.

**LINE GRAPH**

Line graphs illustrate the temporal relationship between two major variables.

**MARKER SELECTION**

Marker Selection is a display of the top differentially expressed genes between two specified cohorts. .

**MESH ONTOLOGY**

MeSH is the National Library of Medicine's controlled vocabulary thesaurus. It consists of sets of terms naming descriptors in a hierarchical structure that permits searching at various levels of specificity.

**MICROARRAY**

A two-dimensional array on a chip or solid surface that assays large amounts of DNA material.

**MRNA ANALYSIS**

Assays that quantify the expression levels of all mRNA molecules in an experiment.

**NAVIGATION TREE**

The Window's Explorer-like, hierarchical representation of study data that has been loaded into Dataset Explorer.

## **NCBI**

The National Center for Biotechnology Information.

See [http:// www.ncbi.nlm.nih.gov/](http://www.ncbi.nlm.nih.gov/) for details.

## **NUMERIC-NODE**

Used in Dataset Explorer, numeric-nodes are indicated by the (123) symbol, numeric nodes indicate that the data values associated with the concept are only numeric (for example, age values, date values, etc.). For more information, see [Continuous Variable](#).

## **ONTOLOGY**

A hierarchical description of the concepts and relationships that can exist for an agent or a community of agents.

## **ORTHOGONAL COMPONENT**

When performing statistical analysis, independent variables that affect a particular dependent variable are said to be orthogonal if they are uncorrelated, since the covariance forms an inner product.

## **PATHOLOGY**

The study of diagnosis and disease.

## **PATHWAY**

A group of genes interacting to form an aggregate biological function.

## **PEARSON CORRELATION**

Obtained by dividing the covariance of the two variables by the product of their standard deviations

## **PRINCIPAL COMPONENT ANALYSIS**

A Principal Component Analysis (PCA) is commonly used as a tool in exploratory data analysis. Data is split into orthogonal components, and the genes/probes that contribute the most variance to the components are displayed.

## **PROBE SET**

A probe set is a collection of probes designed to interrogate a given sequence.

**PROBE SET ID**

A probe set ID is used to refer to a probe set, which looks like the following:

12345\_at or 12345\_a\_at or 12345\_s\_at or 12345\_x\_at

The last three characters (\_at) identify the probe set strand.

**P-VALUE**

The number corresponding probability that the occurrences of your experiment and analysis did not happen by chance. P-value cutoffs are often 0.05 or 0.01 – when the value is under the threshold, the result is said to be statistically significant.

**R**

R is a language and environment for statistical computing and graphics.

See <http://www.r-project.org> for details.

**RBM DATA**

Rules Based Medicine. They provide an array measurement of metabolites

**REGRESSION ALGORITHMS**

Algorithms that are particularly suited for mining data sets that have high dimensionality (many attributes), including transactional and unstructured data.

**RHO-VALUE**

Also known as Spearman's rho, the rho-value is a non-parametric measure of statistical dependence between two variables. See: [Spearman Correlation](#).

**R-VALUE**

The value assigned to a correlation coefficient.

**SAMPLE EXPLORER**

Sample Explorer lets you search for tissue and blood samples of interest so that you can learn more about the samples; for example, you can look up sample IDs and locate the study that produced the samples in the Dataset Explorer.

**SCATTER PLOT**

Type of graph that uses Cartesian coordinates to display values for two variables for a set of data.

**SEARCH FILTER**

A biomedical concept used to define search criteria in the Search tool.

**SEARCH STRING**

A sequence of biomedical concepts used to define search criteria in the Search tool.

**SLOPE**

The steepness of the line of best fit in a graph ( $\Delta y/\Delta x$ ).

**SNP DATA**

Single Nucleotide Polymorphism. DNA sequence data marking variation occurring when a single nucleotide — A, T, C or G — in the genome.

**SPEARMAN CORRELATION**

The Spearman's rank-order correlation is the nonparametric version of the Pearson product-moment correlation. Spearman's correlation coefficient, ( $\rho$ , also signified by rho-value) measures the strength of association between two ranked variables.

**STATISTICAL SIGNIFICANCE**

Results of analyses on data that are statistically significant indicate a confidence level that the results did not happen by chance.

**STUDY GROUP**

The subjects in a study grouped together due to some common attribute of interest (for example, a study can have two study groups: normal and control).

**SUBSET**

A smaller grouping of participants in a study. See [cohort](#).

**SURVIVAL ANALYSIS**

Assessment of the amount of time that a person or population lives after a particular intervention or condition.



**T STATISTIC**

Ratio of the departure of an estimated parameter from its notional value and its standard error.

**TABLE WITH FISHER TEST**

Examines the significance of associated categorical variables.

**TEA ANALYSES**

Target Enrichment Analysis (TEA) measures the enrichment of a gene signature, gene list, or pathway in a microarray expression experiment.

**TEA P-VALUE**

These normalized p-values are intermediate values in the TEA calculation. To be considered a statistically significant analysis, an analysis must have at least one matching biomarker with a TEA p-Value of less than 0.05.

**TEA SCORE****TEXT-NODE**

Indicated by the (abc) symbol, text nodes indicate that the data values associated with the concept are only textual (for example, race or gender). For more information, see [Categorical Variable](#).

**TISSUE TYPE**

The specific type of tissue that has been used in the experiment (for example, breast tissue, lung tissue, etc.)

**UP-REGULATION**

An analysis of a statistically significant experiment returned from a search against a pathway is designated as *up-regulated* or *down-regulated*.

**X-AXIS**

The horizontal axis of a two-dimensional Cartesian coordinate system.

**Y-AXIS**

The vertical axis of a two-dimensional Cartesian coordinate system.

