Package ‘SVMDO’

January 27, 2024

Title Identification of Tumor-Discriminating mRNA Signatures via Support Vector Machines Supported by Disease Ontology

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Date 2024-01-08

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Description It is an easy-to-use GUI using disease information for detecting tumor/normal sample discriminating gene sets from differentially expressed genes. Our approach is based on an iterative algorithm filtering genes with disease ontology enrichment analysis and wilk and wilk’s lambda criterion connected to SVM classification model construction. Along with gene set extraction, SVMDO also provides individual prognostic marker detection. The algorithm is designed for FPKM and RPKM normalized RNA-Seq transcriptome datasets.

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License GPL-3

Encoding UTF-8

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Config/testthat/edition 3

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R topics documented:

classification_server  .................................................. 3
classification_ui ....................................................... 3
clinic_data_input_server .............................................. 4
clinic_data_input_ui .................................................. 4
deg_server ............................................................... 5
deg_ui ................................................................. 5
disc_gene_download_ui .................................................. 6
disc_gene_dw_server .................................................... 6
do_based_gene_filtration_server ...................................... 7
do_based_gene_filtration_ui .......................................... 7
expression_dataset_input_server ..................................... 8
gene_directory_selection_input_server ................................ 8
gene_directory_selection_ui .......................................... 9
gene_list_name_server ................................................. 9
gene_list_name_ui .................................................... 10
gene_list_table_visualization_ui .................................... 10
globals ................................................................. 11
gui_obj_removal_server ............................................... 11
gui_obj_removal_ui .................................................... 12
innerServer_exp_ui ................................................... 12
package_req_list ...................................................... 13
plot_list_server ....................................................... 13
plot_list_ui .......................................................... 14
plot_push_server ...................................................... 14
plot_push_ui .......................................................... 15
plot_show_server ...................................................... 15
plot_show_ui .......................................................... 16
runGUI ................................................................. 16
survival_analysis_server ............................................. 17
survival_analysis_ui .................................................. 17
surv_plot_dw_server ................................................... 18
surv_plot_dw_ui ....................................................... 18
SVMDO ................................................................. 19
table_server .......................................................... 20
classification_server

table_ui .......................... 21
test_data_selection_server .................. 21
test_data_selection_ui ...................... 22
top_val_based_deg_filtration .................. 22
top_val_based_deg_filtration_ui .................. 23
top_val_server .................................. 23
top_val_ui .................................. 24

Index 25

classification_server  SVMDO

Description
SVMDO

Usage
innerServer_7(input, output, session)

Arguments
input server input
output server output
session server session

Value
Server section of wilks lambda filtration and SVM classification of disease filtered differentially expressed gene set

classification_ui  SVMDO

Description
SVMDO

Usage
innerUI_classification(id)

Arguments
id connection input
Value

UI section of wilks lambda filtration and SVM classification of disease filtered differentially expressed gene set

clinic_data_input_server
SVMDO

Description
SVMDO

Usage
innerServer_clinic(input, output, session)

Arguments
input server input
output server output
session server session

Value
Server section of loading clinical data

clinic_data_input_ui
SVMDO

Description
SVMDO

Usage
innerUI_clinic_data(id)

Arguments
id connection input

Value
UI section of loading clinical data
deg_server  

---

deg_server  

SVMD0

Description
SVMD0

Usage
innerServer_3(input, output, session, rawData, rval)

Arguments
input  
server input
output  
server output
session  
server session
rawData  
expression dataset provided from innerServer_exp_server
rval  
Selected radio button information provided from innerServer_rad_server

Value
Server section of differential gene expression analysis

---

deg_ui  

SVMD0

Description
SVMD0

Usage
innerUI_deg_analysis(id)

Arguments
id  
connection input

Value
UI section of differential gene expression analysis
**disc_gene_download_ui**  

**Description**  
SVMDO

**Usage**  

disc_gene_download_ui(id)

**Arguments**

id  
connection input

**Value**  
UI section of discriminative gene set download button

---

**disc_gene_dw_server**  

**Description**  
SVMDO

**Usage**  

disc_gene_dw_server(input, output, session, gene_list_val)

**Arguments**

input  
server input

output  
server output

session  
server session

gene_list_val  
discriminative gene set list variable

**Value**

Server section of discriminative gene set download button
### Description

**SVMDO**

### Usage

`innerServer_6(input, output, session)`

### Arguments

- **input**: server input
- **output**: server output
- **session**: server session

### Value

Server section of disease ontology based filtration of differentially expressed genes

---

### Description

**SVMDO**

### Usage

`innerUI_disease_ont_class(id)`

### Arguments

- **id**: connection input

### Value

UI section of disease ontology based filtration of differentially expressed genes
expression_dataset_input_server
  SVMDO

Description
  SVMDO

Usage
  innerServer_exp(input, output, session)

Arguments
  input  server input
  output server output
  session server session

Value
  Server section of providing expression dataset

gene_directory_selection_server
  SVMDO

Description
  SVMDO

Usage
  innerServer(input, output, session)

Arguments
  input  server input
  output server output
  session server session

Value
  Server section of entering output/working for gene list directory
**gene_directory_selection_ui**

**Description**

SVMDO

**Usage**

innerUI_path(id)

**Arguments**

- id: connection input

**Value**

UI section of entering output/working for gene list directory

---

**gene_list_name_server**

**Description**

SVMDO

**Usage**

innerServer_10(input, output, session)

**Arguments**

- input: server input
- output: server output
- session: server session

**Value**

Server section of entering final gene list name
Description

SVMDO

Usage

gene_list_table_visualization_ui(id)

Arguments

id

Value

Providing table form of discriminative gene sets in GUI
### globals

<table>
<thead>
<tr>
<th>Description</th>
<th>SVMDO</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Value</strong></td>
<td></td>
</tr>
<tr>
<td>Including script files and global variables of GUI required to be initiated at the runApp file execution</td>
<td></td>
</tr>
</tbody>
</table>

### gui_obj_removal_server

<table>
<thead>
<tr>
<th>Description</th>
<th>SVMDO</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Usage</strong></td>
<td></td>
</tr>
<tr>
<td><code>innerServer_9(input, output, session)</code></td>
<td></td>
</tr>
<tr>
<td><strong>Arguments</strong></td>
<td></td>
</tr>
<tr>
<td><code>input</code></td>
<td>server input</td>
</tr>
<tr>
<td><code>output</code></td>
<td>server output</td>
</tr>
<tr>
<td><code>session</code></td>
<td>server session</td>
</tr>
<tr>
<td><strong>Value</strong></td>
<td></td>
</tr>
<tr>
<td>Server section of workspace clearance</td>
<td></td>
</tr>
</tbody>
</table>
**Description**

SVMDO

**Usage**

innerUI_clear_env(id)

**Arguments**

id  connection input

**Value**

UI section of workspace clearance

---

**Description**

SVMDO

**Usage**

innerUI_exp_data(id)

**Arguments**

id  connection input

**Value**

UI section of providing expression dataset into GUI
Description

SVMDO

Value

List of packages involved in SVMDO

---

Description

SVMDO

Usage

plot_list_server(input, output, session)

Arguments

input    server input
output   server output
session  server session

Value

Server section of preparing plot list to be visualized in GUI page
plot_list_ui  SVMDO

Description
SVMD

Usage
innerUI_collect_plot_data(id)

Arguments
id  connection output

Value
UI section of preparing plot list to be visualized in GUI page

plot_push_server  SVMDO

Description
SVMD

Usage
plot_push_server(input, output, session)

Arguments
input  server input
output  server output
session  server session

Value
Server section of providing information about total number of survival plots for visualization
**plot_push_ui**

**Description**

SVMDO

**Usage**

`innerUI_plot_inject(id)`

**Arguments**

- **id**: connection input

**Value**

UI section of providing information about total number of survival plots for visualization

---

**plot_show_server**

**Description**

SVMDO

**Usage**

`plot_show_server(input, output, session, max_data)`

**Arguments**

- **input**: server input
- **output**: server output
- **session**: server session
- **max_data**: Information of total number of survival plots prepared with discriminative gene set

**Value**

Server section of providing information about total number of survival plots for visualization
**plot_show_ui**

### Description

SVMDO

### Usage

innerUI_plot_show(id)

### Arguments

- **id**
  - connection input

### Value

UI section of providing information about total number of survival plots for visualization

---

**runGUI**

### Description

SVMDO

### Usage

linebreaks(n)

### Arguments

- **n**
  - linebreak function variable

### Value

Returning GUI window screen

### Examples

```r
# SVMDO::runGUI() Calling GUI without activating library
# runGUI() Calling GUI after activating library
# Disease Ontology Enrichment of a differentially expressed gene (entrez id):
an_1<-DOSE::enrichDO(2981,ont="DO")
```
survival_analysis_server

\textit{SVMDO}

\textbf{Description}

\textit{SVMDO}

\textbf{Usage}

innerServer_8(input, output, session, rawData_2, rval)

\textbf{Arguments}

\begin{itemize}
  \item \textit{input} server input
  \item \textit{output} server output
  \item \textit{session} server session
  \item \textit{rawData_2} Clinical data provided from clinic_data_input_server
  \item \textit{rval} Selected radio button information provided from innerServer_rad_server
\end{itemize}

\textbf{Value}

Server section of survival analysis of final discriminative gene set

---

survival_analysis_ui \textit{SVMDO}

\textbf{Description}

\textit{SVMDO}

\textbf{Usage}

innerUI_surv(id)

\textbf{Arguments}

\begin{itemize}
  \item \textit{id} connection input
\end{itemize}

\textbf{Value}

UI section of survival analysis of final discriminative gene set
surv_plot_dw_server  SVMDO

Description

SVMD

Usage

surv_plot_dw_server(input, output, session)

Arguments

input  server input
output  server output
session  server session

Value

Server section of downloading survival plots of discriminative gene set

---

surv_plot_dw_ui  SVMDO

Description

SVMD

Usage

surv_plots_download_ui(id)

Arguments

id  connection input

Value

UI section of downloading survival plots of discriminative gene set
Description

Package Description: It is an easy-to-use GUI using disease information for detecting tumor/normal sample discriminating gene sets from differentially expressed genes. Our approach is based on an iterative algorithm filtering genes with disease ontology enrichment analysis and wilk’s lambda criterion connected to SVM classification model construction. Along with gene set extraction, SVMDO also provides individual prognostic marker detection among the discriminative genes. The algorithm is designed for FPKM and RPKM normalized RNA-Seq transcriptome datasets. To provide experience about the GUI usage, a test section involving dummy example using SummarizedExperiment objects of transcriptome (small form) and clinical datasets is also included.

Value

Providing package-level manual page

Package Sections

1. Analysis: Acquiring discriminative gene sets and further detecting the gene subset with prognostic characteristics
2. Result: Visualization and Download of discriminative gene sets and survival plot list of prognostic genes

Steps of Analysis Screen

1. To search your transcriptome dataset, use the file detection in Choose Your Expression Dataset section. The file will be automatically uploaded into the GUI.
2. To prevent clashing with test datasets, "None" option has to be selected from the radio button section.
3. By clicking on DEG Analysis button you further apply differential expression analysis. Labels of tissue_type column in dataset must contain “Nor” and “Tum” for determining normal/tumour (or tumor) samples. A message window saying Process Completed will appear if there is not any problem.
4. When the differential expression process is completed, a user-defined input size (n) is selected to filter the initial gene list (i.e., n number of upregulated and downregulated genes) by entering a number in Input Size section. It is predetermined as 50 in GUI which can be changed based on the user. If there is problem with the value of input size, you will get a warning about inappropriate input size selection. If the input size remains, algorithm selects all of the differentially expressed genes to be used in the next process.
5. To apply disease ontology-based gene filtration, click on DO Analysis button. A message window saying process completed will appear if there is not any problem.
6. To further apply the following feature selection and classification processes, click on the Classification button. A message window saying process completed will appear if there is not any problem.
7. Acquired discriminative gene set can be further used for survival analysis to detect individual prognostic genes. To apply this process, use the file detection in Choose Clinical Data section for searching clinical data about patient survival followed by clicking on Survival Analysis button.

Steps of Result Screen

1. To visualize discriminative gene sets inside GUI screen, click on Show Gene Results button. When you click this button, a table of gene set will appear. If there is a problem in the analysis, an error message will appear.

2. To visualize survival plots of individual genes, two steps have to be applied. First of all, click on Prepare Plot Lists button to feed plot information to the visualization system. After that, click on Show Plots button to visualize survival plots.

3. Before downloading files, you can adjust the output directory with Choose Directory button. It can be used for separating files by selecting a destination before clicking download buttons. If it is desired, files can be downloaded to the same folder by selecting an output directory just one time before the download steps. If you do not select any output directory, files will be downloaded to your working directory.

4. To download the resulting discriminative gene set, it is obligatory to define a filename in the Enter Final Gene Set Filename section. After that, you can click on Download Gene List button to complete the process.

5. To download survival plots, you have to click on Download Plot List button. Names of plot files are automatically done by assigning gene names.

Application of Test Datasets

SVMDO includes test datasets providing dummy examples for gaining experience on the GUI usage. Test datasets consist of simplified forms of TCGA-COAD (COAD) and TCGA-LUSC (LUSC) with 400 genes along with clinical datasets loaded into summarized experiment objects. When test datasets are used, predetermined expression and clinical datasets are automatically uploaded into the GUI. A test-based analysis is done with predefined input size (n=50). Therefore, users have to continue with DO Analysis after DEG Analysis.

Workspace Clearance

When the user task is completed, click on the Clear Environment button to remove the global variables created during the algorithm sections. To prevent error in the next usages of GUI, it is a necessary process. It can be applied at any moment without the necessity of completing all of the steps of algorithm.

---

**Description**

SVMDO
**Usage**

```
table_server(input, output, session)
```

**Arguments**

- **input**: server input
- **output**: server output
- **session**: server session

**Value**

Server section of providing discriminative gene set for preparing table

---

**Description**

SVMDO

**Usage**

```
innerUI_table_show(id)
```

**Arguments**

- **id**: connection input

**Value**

UI section of providing discriminative gene set for preparing table

---

**Description**

SVMDO

**Usage**

```
innerServer_rad(input, output, session)
```
Arguments

<table>
<thead>
<tr>
<th>Argument</th>
<th>Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>input</td>
<td>server input</td>
</tr>
<tr>
<td>output</td>
<td>server output</td>
</tr>
<tr>
<td>session</td>
<td>server session</td>
</tr>
</tbody>
</table>

Value

Server section of providing information about selected radio button

---

test_data_selection_ui

\[ SVMDO \]

Description

SVMDO

Usage

innerUI_test_data(id)

Arguments

<table>
<thead>
<tr>
<th>Argument</th>
<th>Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>id</td>
<td>connection input</td>
</tr>
</tbody>
</table>

Value

UI section of providing information about selected radio button

---

top_val_based_deg_filtration

\[ SVMDO \]

Description

SVMDO

Usage

innerServer_5(input, output, session, top_val)
**Arguments**

- input: server input
- output: server output
- session: server session
- top_val: top gene number value provided from top_val_server

**Value**

Server section of selecting differentially expressed genes based on top gene value

---

**Description**

SVMDO

**Usage**

`innerUI_top_gene_selection(id)`

**Arguments**

- id: connection input

**Value**

UI section of selecting differentially expressed genes based on top gene value

---

**Description**

SVMDO

**Usage**

`innerServer_4(input, output, session)`

**Arguments**

- input: server input
- output: server output
- session: server session
Value

Server section of entering top gene value

---

top_val_ui  SVMDO

Description

SVMDO

Usage

innerUI_top_gene_val(id)

Arguments

id  connection input

Value

UI section of entering top gene value
Index

classification_server, 3
classification_ui, 3
clinic_data_input_server, 4
clinic_data_input_ui, 4
deg_data_table_ui
   (gene_list_table_visualization_ui), 10
deg_server, 5
deg_ui, 5
disc_gene_download_ui, 6
disc_gene_dw_server, 6
do_based_gene_filtration_server, 7
do_based_gene_filtration_ui, 7
expression_dataset_input_server, 8
gene_directory_selection_server, 8
gene_directory_selection_ui, 9
gene_list_name_server, 9
gene_list_name_ui, 10
gene_list_table_visualization_ui, 10
globals, 11
gui_obj_removal_server, 11
gui_obj_removal_ui, 12
innerServer
   (gene_directory_selection_server), 8
   innerServer_10 (gene_list_name_server), 9
   innerServer_3 (deg_server), 5
   innerServer_4 (top_val_server), 23
   innerServer_5
      (top_val_based_deg_filtration), 22
   innerServer_6
      (do_based_gene_filtration_server), 7
   innerServer_7 (classification_server), 3
   innerServer_8
      (survival_analysis_server), 17
   innerServer_9 (gui_obj_removal_server), 11
innerServer_clinic
   (clinic_data_input_server), 4
innerServer_exp
   (expression_dataset_input_server), 8
   innerServer_exp_ui, 12
innerServer_rad
   (test_data_selection_server), 21
innerUI_classification
   (classification_ui), 3
innerUI_clear_env (gui_obj_removal_ui), 12
innerUI_clinic_data
   (clinical_data_input_ui), 4
innerUI_collect_plot_data
   (plot_list_ui), 14
innerUI_deg_analysis (deg_ui), 5
innerUI_disease_ont_class
   (do_based_gene_filtration_ui), 7
innerUI_exp_data (innerServer_exp_ui), 12
innerUI_gene_names (gene_list_name_ui), 10
innerUI_path
   (gene_directory_selection_ui), 9
innerUI_plot_inject (plot_push_ui), 15
innerUI_plot_show (plot_show_ui), 16
innerUI_surv (survival_analysis_ui), 17
innerUI_table_show (table_ui), 21
innerUI_test_data
   (test_data_selection_ui), 22
innerUI_top_gene_selection
(top_val_based_deg_filtration_ui), 23
innerUI_top_gene_val(top_val_ui), 24

linebreaks (runGUI), 16

package_req_list, 13
plot_list_server, 13
plot_list_ui, 14
plot_push_server, 14
plot_push_ui, 15
plot_show_server, 15
plot_show_ui, 16

runGUI, 16

surv_plot_dw_server, 18
surv_plot_dw_ui, 18
surv_plots_download_ui
  (surv_plot_dw_ui), 18
survival_analysis_server, 17
survival_analysis_ui, 17
SVMDO, 19

table_server, 20
table_ui, 21
test_data_selection_server, 21
test_data_selection_ui, 22
top_val_based_deg_filtration, 22
top_val_based_deg_filtration_ui, 23
top_val_server, 23
top_val_ui, 24