# Package 'CELLector'

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Type Package

Title Genomics guided selection of cancer cell lines
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<b>Description</b> Functions to select the most relevant cancer cell lines to be included in a new invitro study, in a genomic-guided fashion. CELLector combines methods from graph theory and market basket analysis; it leverages tumour genomics data to explore, rank, and select optimal cell line models in a user-friendly way, enabling scientists to make appropriate and informed choices about model inclusion/exclusion in retrospective analyses and future studies. Additionally, it allows the selection of models within user-defined contexts, for example, by focusing on genomic alterations occurring in biological pathways of interest or considering only predetermined sub-cohorts of cancer patients. Finally, CELLector identifies combinations of molecular alterations underlying disease subtypes currently lacking representative cell lines, providing guidance for the future development of new cancer models.
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Encoding UTF-8
LazyData true
<b>Depends</b> arules, dplyr, stringr, data.tree, sunburstR, igraph, collapsibleTree, methods, RCurl
Suggests knitr, rmarkdown
VignetteBuilder knitr
R topics documented:
CELLector.buildModelMatrix CELLector.Build_Search_Space CELLector.CellLine.BEMs CELLector.CFEs CELLector.CFEs CELLector.CFEs.CNAid_decode CELLector.CFEs.CNAid_mapping CELLector.changeSScolors

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CELLector.buildModelMatrix

Mapping cell lines on the CELLector searching space

### **Description**

This function maps cell line on the subtypes identified and assembled in the CELLector searching space, based on the collective presence/absence of the signatures of cancer functional events underlying these subtypes. The subtypes lacking representative cell lines are not considered and, in the output, the subtypes (indicated by their numerical id, which matches that in the CELLector searching space) are ranked based on the greedy algorithm described in [1] based on their covered genomic heterogeneity.

### Usage

CELLector.buildModelMatrix(Sigs, dataset, searchSpace)

### Arguments

Sigs	A vector of string, in which each element represents a signature of cancer functional events (CFEs, defined in [2]) corresponding to a node in the CELLector searching space. This is expressed as a logic formula (rule), which a cancer patient's genome must satisfy in order to be included in the sub-population represented by the node under consideration. This vector is outputted by the CELLector.createAllSignatures function starting from a CELLector searching space (created by the CELLector.Build_Search_Space) function
dataset	A data frame in which the first two columns contain the COSMIC [3] identiefiers and names of cell lines (one per row), respectively, and then binary entries indicating the status of each CFEs (one per column) across cell lines. The format is the same of the entries of the list in the built-in CELLector.CellLine.BEMs object.
searchSpace	A CELLector searching space encoded as binary tree in a navigable table, as

### Value

A named binary matrix with suptypes numerical identifiers on the rows, cell line names on the column and entries specifying whether the cell line in the column is representative of the subtype on the row (based on the collective presence/absence of the corresponding signature of CFEs)

returned by the CELLector.Build\_Search\_Space function

#### Author(s)

Hanna Najgebauer and Francesco Iorio

#### References

[1] Najgebauer, H. et al. Genomics Guided Selection of Cancer in vitro Models. https://doi.org/10.1101/275032

- [2] Iorio, F. et al. A Landscape of Pharmacogenomic Interactions in Cancer. Cell 166, 740–754 (2016).
- [3] Forbes, S. A. et al. COSMIC: exploring the world's knowledge of somatic mutations in human cancer. Nucleic Acids Res. 43, D805–11 (2015).

### See Also

```
CELLector.CreateAllSignatures,
CELLector.Build_Search_Space,
CELLector.CellLine.BEMs,
CELLector.Build_Search_Space
```

```
data(CELLector.PrimTum.BEMs)
data(CELLector.Pathway_CFEs)
data(CELLector.CFEs.CNAid_mapping)
data(CELLector.CFEs.CNAid_decode)
data(CELLector.HCCancerDrivers)
data(CELLector.CellLine.BEMs)
### Change the following two lines to work with a different cancer type
tumours_BEM<-CELLector.PrimTum.BEMs$COREAD</pre>
CELLlineData<-CELLector.CellLine.BEMs$COREAD
### unicize the sample identifiers for the tumour data
tumours_BEM<-CELLector.unicizeSamples(tumours_BEM)</pre>
### building a CELLector searching space focusing on three pathways
### and TP53 wild-type patients only
CSS<-CELLector.Build_Search_Space(ctumours = t(tumours_BEM),</pre>
                                   verbose = FALSE,
                                   minGlobSupp = 0.05,
                                   cancerType = 'COREAD',
                                   pathwayFocused = c("RAS-RAF-MEK-ERK / JNK signaling",
                                                      "PI3K-AKT-MTOR signaling",
                                                      "WNT signaling"),
                                   pathway_CFEs = CELLector.Pathway_CFEs,
                                   cnaIdMap = CELLector.CFEs.CNAid_mapping,
                                   cnaIdDecode = CELLector.CFEs.CNAid_decode,
                                   cdg = CELLector.HCCancerDrivers,
                                   subCohortDefinition='TP53',
                                   NegativeDefinition=TRUE)
```

```
### take all the signatures from the searching space
Signatures <- CELLector.createAllSignatures(CSS$navTable)

### mapping colorectal cancer cell lines onto the CELLector searching space
ModelMat<-CELLector.buildModelMatrix(Signatures$ES,CELLlineData,CSS$navTable)
head(ModelMat)</pre>
```

```
CELLector.Build_Search_Space
```

CELLector search space construction

### **Description**

This function assembles a user defined CELLector search space analysing genomic data from a larg cohort of cancer patients (spcified in input). It identifies recurrent subtypes with matched genomic signatures (as combination of cancer functional events (CFEs), defined in [1]), linking them into a hierarchical structure shaped as a binary three with a corresponding navigable table, as detailed in [2].

### Usage

### **Arguments**

ctumours	A binary event matrix (BEM) modeling a cohort of cancer patients. With cancer functional events (CFEs) on the columns and sample identifiers on the rows. See CELLector.PrimTum.BEMs for further details
cancerType	The cancer type under consideration (speified via a TCGA label): currently available types = <i>BLCA</i> , <i>BRCA</i> , <i>COREAD</i> , <i>GBM</i> , <i>HNSC</i> , <i>KIRC</i> , <i>LAML</i> , <i>LGG</i> , <i>LUAD</i> , <i>LUSC</i> , <i>OV</i> , <i>PRAD</i> , <i>SKCM</i> , <i>STAD</i> , <i>THCA</i> , <i>UCEC</i>
minlen	The minimal length of the genomic signatures (how many indivudal CFEs it is made of) in order to be considered in the analysis (1 by default)
verbose	A boolean argument specifying whether step-by-step information on the algorithm progression should be displayed run-time

mutOnly A boolean argument specifying whether only CFEs involving somatic mutations

should be considered in the analysis. If the cnaOnly argument is equal to TRUE

then this must be FALSE (default value)

cnaOnly A boolean argument specifying whether only CFEs involving copy number alter-

ations (CNAs) of chromosomal segments that are recurrently CN altered should be considered in the analysis. If the mutOnly argument is equal to TRUE then this

must be FALSE (default value)

minGlobSupp Minimal size of the outpputted subtypes, as ratio of the number patients included

in the whole cohort.

FeatureToExclude

A string (or a vector of strings) with identifiers of CFEs that should be ignored

pathway\_CFEs A named list of string vectors, whose elements are CFEs involving genes in a biological pathway (specified by the name of the corresponding entry). A list for 14 key cancer pathways is contained in the CELLector.Pathway\_CFEs data

object (see corresponding help page for further deatails)

pathwayFocused If different from NULL (default value), it should be a vector of strings. In this

case the analysis will consider only CFEs involving genes in a set of pathways, whose names are contained in this argument and must be present as names of

the  $pathway\_CFEs$  argument

subCohortDefinition

If different from NULL (default value), it should be a string containing the identifier of a CFE. In this case the analysis will consider only the primary tumour samples harbouring (or not harbouring, depending on the NegativeDefinition

argument) the specified CFE

NegativeDefinition

If the subCohortDefinition argument is not NULL then this paramenter determines whether to consider primary tumour samples that harbour (if equal to

FALSE, default value) or not (if equal to TRUE) the specified CFE

cnaIdMap A data frame mapping chromosomal regions of recurrent copy number amplifi-

cations/deletions in cancer (RACSs, as defined in [1]) identified via ADMIRE [3] in the context of specific cancer types to PanCancer RACSs. The built-in object CELLector.CFEs.CNAid\_mapping (or an alternative data frame with the

same format) should be used.

cnaIdDecode A table with identifiers of cancer functional events (CFEs) involving chromo-

somal regions of recurrent copy number alterations (RACSs, as defined by [1], i.e. identified throgh ADMIRE [3]) and their annotation. The built-in object CELLector.CFEs.CNAid\_decode (or an alternative data frame with the same

format) should be used.

A list of genes that are used when decoding the identifiers of cancer functional events (CFEs) involving chromosomal regions of recurrent copy num-

ber alterations (RACSs, as defined by [1]). These will be visualised in the signatures containing the RACSs including them. A predefined list of high confidence cancer driver genes (from [1]) is provided as built-in data object

(CELLector.HCCancerDrivers)

### **Details**

Starting from an initial cohort of patients affected by a given cancer type and modeled by the inputted binary event matrix (BEM), the most frequent alteration or set of molecular alterations (depending on the minlen argument) with the largest support (the subpopulation of patients in

which these alterations occur simultaneously) is identified using the eclat function of the arules R package.

Based on this, the cohort of patients is split into two subpopulations depending on the collective presence or absence of the identified alterations. This process is then executed recursively on the two resulting subpopulations and it continues until all the alteration sets (with a support of minimal size, as specified in the minGlobSupp argument) are identified.

Each of the alterations sets identified through this recursive process is stored in a tree node. Linking nodes identified in adjacent recursions yields a binary tree: the CELLector search space. Each individual path (from the root to a node) of this tree defines a rule (signature), represented as a logic AND of multiple terms (or their negation), one per each node in the path. If the genome of a given patient in the analysed cohort satisfies the rule then it is contained in the subpopulation represented by the terminal node of that path. Collectively, all the paths in the search space provide a representation of the spectrum of combinations of molecular alterations observed in a given cancer type, and their clinical prevalence in the analysed patient population.

#### Value

A named list with the CELLector search space stored as a data.tree object in the TreeRoot field and as a *navigable table*: a data frame with a row for each node of the tree and the following columns

Idx A numerical index for the node

Item The most supported CFE (or a combination of CFE), identified at the iteration in which the node has been added to the three, (i) in the whole cohort of patients (for the Root), (ii) in the sub population that satisfies the parent node rule (for Left.Child nodes) or (iii) its complement (for Right.Child nodes)

ItemsDecoded Same as Item but with identifiers of RACSs decoded, i.e. with loci and included driver genes (inputted in the cdg argument), indicated among brackets

Type The node type: Root (first node added), Right.Child (a node resulting from the analyses of the complementar population of patients with respect to that satisfifying the Parent node rule), Left.Child (a node resulting from refining the population of patients satisfifying the Parent node rule)

Parent.Idx The numerical index of the parent node (0 for the Root)

AbsSupport The number of patients satisfying the node rule

CurrentTotal The number of patients included in the population under consideration at the iteration time of the node inclusion in the tree, this is the same of the parent's AbsSupport for Left.Child nodes

PercSupport The ratio of patients collectively harbouring the combination of CFEs specified in Items within the subpopulation under consideration at the iteration time of the node inclusion in the tree (whose size is specified in CurrentTotal)

GlobalSupport The ratio of patients satisfying the node rule with respect to the total number of patients in the whole cohort

Left.Child.Index Numerical index of the left child node (0 indicates absence of a left child node)

Right.Child.Index Numerical index of the right child node (0 indicates absence of a right child node)

currentPoints The identifiers of the patients in the sub-population under consideration at the iteration time of the node inclusion in the tree

currentFeatures The CFEs considered at the at the iteration time of the node inclusion in the tree positivePoints The identifiers of the patients satisfying the node rule

COLORS A vector of strings containing hexadecimal color identifiers: one for each node. These are used by the visualisation functions (CELLector.visualiseSearchingSpace, and CELLector.visualiseSearchingSpace, and

#### Author(s)

Hanna Najgebauer and Francesco Iorio

### References

- [1] Iorio, F. et al. A Landscape of Pharmacogenomic Interactions in Cancer. Cell 166, 740–754 (2016).
- [2] Najgebauer, H. et al. Genomics Guided Selection of Cancer in vitro Models.

https://doi.org/10.1101/275032

[3] van Dyk, E., Reinders, M. J. T. & Wessels, L. F. A. A scale-space method for detecting recurrent DNA copy number changes with analytical false discovery rate control. Nucleic Acids Res. 41, e100 (2013).

#### See Also

```
CELLector.PrimTum.BEMs,
CELLector.Pathway_CFEs,
CELLector.CFEs.CNAid_mapping,
CELLector.CFEs.CNAid_decode,
CELLector.HCCancerDrivers,
CELLector.visualiseSearchingSpace,
CELLector.visualiseSearchingSpace_sunBurst,
CELLector.changeSScolors
```

```
data(CELLector.PrimTum.BEMs)
data(CELLector.Pathway_CFEs)
data(CELLector.CFEs.CNAid_mapping)
data(CELLector.CFEs.CNAid_decode)
data(CELLector.HCCancerDrivers)
data(CELLector.CellLine.BEMs)
### Change the following two lines to work with a different cancer type
tumours_BEM<-CELLector.PrimTum.BEMs$COREAD</pre>
CELLlineData<-CELLector.CellLine.BEMs$COREAD
### unicize the sample identifiers for the tumour data
tumours_BEM<-CELLector.unicizeSamples(tumours_BEM)</pre>
### building a CELLector searching space focusing on three pathways
### and TP53 wild-type patients only
CSS<-CELLector.Build_Search_Space(ctumours = t(tumours_BEM),</pre>
                                   verbose = FALSE,
                                   minGlobSupp = 0.05,
```

### visualising the CELLector searching space as a binary tree
CSS\$TreeRoot

```
### visualising the first attributes of the tree nodes
CSS$navTable[,1:11]
```

### visualising the sub-cohort of patients whose genome satisfies the rule of the 4th node
str\_split(CSS\$navTable\$positivePoints[4],',')

CELLector.CellLine.BEMs

Cell Lines' Binary Event Matrices

### **Description**

A list containing 16 data frames (one for cancer type), identified through TCGA labels. Each of these data frames contains cell lines' *binary event matrices* (BEMs) with the status (presence/absence) of *cancer functional events* (CFEs) as defined in [1].

### Usage

```
data(CELLector.CellLine.BEMs)
```

### **Format**

A named list of data frames (with TCGA cancer type labels as names). Each of these data frames contains two columns with COSMIC [2] identiefiers and names of cell lines (one per row), respectively, and then binary entries indicating the status of each CFEs (one per column) across cell lines.

### Details

BEMs for cell lines from the Genomics of Drug Sensitivity in Cancer (GDSC1000, [1]) panel. Data is available for cell lines matching one among 16 different TCGA cancer types: *BLCA, BRCA, COREAD, GBM, HNSC, KIRC, LAML, LGG, LUAD, LUSC, OV, PRAD, SKCM, STAD, THCA, UCEC*.

A decoding table for these labels is available at

https://www.cancerrxgene.org/gdsc1000/GDSC1000\_WebResources/Data\_Summary.html.

Each data frame contains cell lines on the rows (with COSMIC identifiers and names, respectively on first and second column) and then a binary matrix with a CFE per column and entries indicating the presence/absence of a given CFE in a given cell line.

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Gene symbols as column names indicate high confidence cancer driver genes and the entries in the corresponding columns indicate the presence/absence of somatic mutations. Column names with *cna* as prefix indicate chromosomal segments that are recurrently copy number altered in cancer (RACSs, defined in [1]). A list with all the considered CFEs is available in the CELLector.CFEs data object. A decoding table for the RACSs is available in the CELLector.CFEs.CNAid\_decode, with the mapping realised by the values in the CNA\_identifier column.

Please note that the same RACS identifier across multiple cancer types might indicate different chromosomal regions, therefore in order to be decode it should be considered jointly with the TCGA label of the data frame it has been extracted from.

### Author(s)

Francesco Iorio (fi9232@gmail.com)

#### References

- [1] Iorio, F. et al. A Landscape of Pharmacogenomic Interactions in Cancer. Cell 166, 740–754 (2016).
- [2] Forbes, S. A. et al. COSMIC: exploring the world's knowledge of somatic mutations in human cancer. Nucleic Acids Res. 43, D805–11 (2015).

### See Also

```
{\tt CELLector.PrimTum.BEMs, CELLector.CFEs, CELLector.CFEs.CNAid\_decode}
```

### **Examples**

CELLector.CFEs

Cancer Functional Events

### **Description**

Identifiers of cancer functional events (CFEs, i.e. somatic mutations in high confidence cancer driver genes or chromosomal regions of recurrent copy number amplification/deletion) from [1], which are also present in the binary event matrices of the cell lines and the primary tumours considered in this version of CELLector.

### Usage

```
data("CELLector.CFEs")
```

### **Format**

A vector of strings with one entry per identifier.

#### **Details**

Gene symbols indicate somatic mutations in igh confidence cancer driver genes and entries with *cna* prefix indicate chromosomal segments that are recurrently copy number altered in cancer (RACSs), both defined in [1].

A decoding table for the RACSs is available in the CELLector. CFEs. CNAid\_decode, with the mapping realised by the values in the CNA\_identifier column.

Please note that the same RACS identifier across multiple cancer types might indicate different chromosomal regions, therefore in order to be decode it should be considered jointly with the TCGA label of the data frame it has been extracted from.

#### Author(s)

Francesco Iorio (fi9232@gmail.com)

#### References

[1] Iorio, F. et al. A Landscape of Pharmacogenomic Interactions in Cancer. Cell 166, 740–754 (2016).

### See Also

CELLector.PrimTum.BEMs, CELLector.CellLine.BEMs, CELLector.CFEs, CELLector.CFEs.CNAid\_decode

#### **Examples**

```
data(CELLector.CFEs)
head(CELLector.CFEs)
```

CELLector.CFEs.CNAid\_decode

Decoding table for copy number alteration cancer functional events

### Description

A table with identifiers of cancer functional events (CFEs) involving chromosomal regions of recurrent copy number alterations (RACSs, as defined by [1], i.e. identified through ADMIRE [2]) and their annotation.

### Usage

```
data("CELLector.CFEs.CNAid_decode")
```

#### **Format**

A data frame with 731 observations (one for each CNA CFE) on the following 15 variables.

Identifier The RACS identifer, as defined in [1]

CancerType A TCGA label indicating the cancer type where the RACS has been identified (via ADMIRE [2])

Recurrent A string specifying whether the RACS under consideration is frequently amplified (value = Amplification) or deleted ) (value = deleted)

chr Chromosome number of the RACS

start Starting position of the RACS

stop Ending position of the RACS

nGenes Number of protein coding genes included in the RACS

locus Genomic locus of the RACS

ContainedGenes A string with comma separated symbols of the genes included in the RACS

CNA\_Identifier A string containing the identifier of the RACS as it appears in the Binary Event Matrix (BEM) of the cancer type specified in the CancerType field included in the CELLector.CellLine.BEMs and the CELLector.PrimTum.BEMs data objects

#### **Details**

This data frame contains a comprehensive annotation of the CFEs involving RACSs appearing in the BEMs of cell lines and primary tumours, contained in the CELLector.CellLine.BEMs and the CELLector.PrimTum.BEMs data objects. Please note that the same RACS identifier across multiple cancer types might indicate different chromosomal regions, therefore in order to be decode it should be considered jointly with the TCGA label of the data frame it has been extracted from.

This table is used by the CELLector.cna\_look\_up function to decode the identifier of CFE involving a RACS.

### Author(s)

Francesco Iorio (fi9232@gmail.com)

### References

[1] Iorio, F. et al. A Landscape of Pharmacogenomic Interactions in Cancer. Cell 166, 740–754 (2016).

[2] van Dyk, E., Reinders, M. J. T. & Wessels, L. F. A. A scale-space method for detecting recurrent DNA copy number changes with analytical false discovery rate control. Nucleic Acids Res. 41, e100 (2013).

### See Also

```
CELLector.CellLine.BEMs, CELLector.PrimTum.BEMs, CELLector.cna_look_up
```

CELLector.CFEs.CNAid\_mapping

Pan-Cancer/Cancer-Specific RACSs map.

### **Description**

A data frame mapping chromosomal regions of recurrent copy number amplifications/deletions in cancer (RACSs, as defined in [1]) identified via ADMIRE [2] in the context of specific cancer types to PanCancer RACSs.

### Usage

```
data("CELLector.CFEs.CNAid_mapping")
```

### **Format**

A data frame with 425 observations (one for each PanCancer RACS) and a column for each of 27 different cancer types (specified by TCGA labels). The entry in position *i,j* contains the identifier of the *i*th PanCancer RACS in the context of the *j*th cancer type (where available).

### Author(s)

Francesco Iorio (fi9232@gmail.com)

### References

- [1] Iorio, F. et al. A Landscape of Pharmacogenomic Interactions in Cancer. Cell 166, 740–754 (2016).
- [2] van Dyk, E., Reinders, M. J. T. & Wessels, L. F. A. A scale-space method for detecting recurrent DNA copy number changes with analytical false discovery rate control. Nucleic Acids Res. 41, e100 (2013).

### **Examples**

```
data(CELLector.CFEs.CNAid_mapping)
head(CELLector.CFEs.CNAid_mapping)
```

CELLector.changeSScolors

Changing CELLector searching space color scheme

### **Description**

This function reassigns a set of distinc random colors to the cancer patients' subtypes identified and stored in the CELLector searching space (compute by the CELLector.Build\_Search\_Space function). These colors are used by the visualisation functions CELLector.visualiseSearchingSpace and CELLector.visualiseSearchingSpace\_sunBurst

#### Usage

```
CELLector.changeSScolors(searchSpace)
```

### **Arguments**

searchSpace A CELLector searching space as returned by the CELLector.Build\_Search\_Space function

### Value

A CELLector searching space as returned by the CELLector.Build\_Search\_Space function, with new colors

#### Author(s)

Hanna Najgebauer and Francesco Iorio

### See Also

```
CELLector.Build_Search_Space,
CELLector.Build_Search_Space,
CELLector.visualiseSearchingSpace_sunBurst
```

```
data(CELLector.PrimTum.BEMs)
data(CELLector.Pathway_CFEs)
data(CELLector.CFEs.CNAid_mapping)
data(CELLector.CFEs.CNAid_decode)
data(CELLector.HCCancerDrivers)
data(CELLector.CellLine.BEMs)
tumours_BEM<-CELLector.PrimTum.BEMs$COREAD
CELLlineData<-CELLector.CellLine.BEMs$COREAD
### unicize the sample identifiers for the tumour data
tumours_BEM<-CELLector.unicizeSamples(tumours_BEM)</pre>
### building the CELLector searching space
CSS<-CELLector.Build_Search_Space(ctumours = t(tumours_BEM),</pre>
                                   verbose = FALSE,
                                   minGlobSupp = 0.05,
                                   cancerType = 'COREAD',
                                   pathway_CFEs = CELLector.Pathway_CFEs,
                                   cnaIdMap = CELLector.CFEs.CNAid_mapping,
                                   mutOnly = FALSE,
                                   cnaIdDecode = CELLector.CFEs.CNAid_decode,
                                   cdg = CELLector.HCCancerDrivers)
### visualising the Searching space as collapsible tree
CELLector.visualiseSearchingSpace(searchSpace = CSS,CLdata = CELLlineData)
### visualising the Searching space as sunburst
CELLector.visualiseSearchingSpace_sunBurst(searchSpace = CSS)
```

```
### changing the searching space colors
CSS<-CELLector.changeSScolors(CSS)

### visualising the Searching space as collapsible tree
CELLector.visualiseSearchingSpace(searchSpace = CSS,CLdata = CELLlineData)
### visualising the Searching space as sunburst
CELLector.visualiseSearchingSpace_sunBurst(searchSpace = CSS)</pre>
```

### **Description**

This functions shows the annotation for a chromosomal region of recurrent copy number alterations (RACS) as defined in [1].

### Usage

```
CELLector.cna_look_up(cna_ID, cnaId_decode, TCGALabel)
```

### Arguments

cna\_ID A string containin the RACS identifier. Full list available in the CELLector. CFEs

object.

cnaId\_decode A data frame containing the RACSs' annotation, available in the

CELLector.CFEs.CNAid\_decode object

TCGALabel A TCGA label indicating the cancer type under consideration: BLCA, BRCA,

COREAD, GBM, HNSC, KIRC, LAML, LGG, LUAD, LUSC, OV, PRAD, SKCM,

STAD, THCA, UCEC available in this version.

#### Value

A data frame with a single line containing the annotation of the RACS indicated in input.

### Author(s)

Hanna Najgebauer and Francesco Iorio

### References

[1] Iorio, F. et al. A Landscape of Pharmacogenomic Interactions in Cancer. Cell 166, 740–754 (2016).

### See Also

```
CELLector.CFEs,
CELLector.CFEs.CNAid_decode
```

CELLector.createAllSignatures

Derive signatures underlying the cancer patients' subtypes in the CELLector search space

### **Description**

This function takes in input the CELLector search space encoded as a binary tree in a *navigable table*. Then, for each individual path (from the root to a node) of this tree it derives a rule (signature), represented as a logic AND of multiple terms (or their negation), one per each node in the path. Negations are added when right branches are encountered.

### Usage

CELLector.createAllSignatures(NavTab)

### **Arguments**

NavTab

A CELLector searching space encoded as binary tree in a navigable table, as returned by the CELLector.Build\_Search\_Space function.

#### Value

A list with two vectors of strings and a numerical vector. Each element of the first two vectors represent a signature of cancer functional events (CFEs, defined in [1]) corresponding to a node in the CELLector searching space. This is expressed as a logic formula (rule), which a cancer patient's genome must satisfy in order to be included in the sub-population represented by the node under consideration. The first vector (S) contains decoded signatures, i.e. where the CFEs involving copy number alterations are represented by a genomic loci and contained cancer driver genes. The second vector (ES) contains signatures of CFEs as they are represented in the binary event matrix containing the patients genomic data used to build the CELLector searching space. Further deatils are provided in [2]. The third vector (STS) contains the percentage of cancer patients belonging to the subtype represented by the signatures.

#### Author(s)

Hanna Najgebauer and Francesco Iorio

#### References

[1] Iorio, F. et al. A Landscape of Pharmacogenomic Interactions in Cancer. Cell 166, 740–754 (2016).

[2] Najgebauer, H. et al. Genomics Guided Selection of Cancer in vitro Models.

https://doi.org/10.1101/275032

### See Also

CELLector.Build\_Search\_Space

#### **Examples**

```
data(CELLector.PrimTum.BEMs)
data(CELLector.Pathway_CFEs)
data(CELLector.CFEs.CNAid_mapping)
data(CELLector.CFEs.CNAid_decode)
data(CELLector.HCCancerDrivers)
data(CELLector.CellLine.BEMs)
### Change the following two lines to work with a different cancer type
tumours_BEM<-CELLector.PrimTum.BEMs$COREAD</pre>
CELLlineData<-CELLector.CellLine.BEMs$COREAD
### unicize the sample identifiers for the tumour data
tumours_BEM<-CELLector.unicizeSamples(tumours_BEM)</pre>
### building a CELLector searching space focusing on three pathways
### and TP53 wild-type patients only
CSS<-CELLector.Build_Search_Space(ctumours = t(tumours_BEM),</pre>
                                   verbose = FALSE,
                                   minGlobSupp = 0.05,
                                   cancerType = 'COREAD',
                                   pathwayFocused = c("RAS-RAF-MEK-ERK / JNK signaling",
                                                       "PI3K-AKT-MTOR signaling",
                                                       "WNT signaling"),
                                   pathway_CFEs = CELLector.Pathway_CFEs,
                                   cnaIdMap = CELLector.CFEs.CNAid_mapping,
                                   cnaIdDecode = CELLector.CFEs.CNAid_decode,
                                   cdg = CELLector.HCCancerDrivers,
                                   subCohortDefinition='TP53',
                                   NegativeDefinition=TRUE)
### derive signatures from searching space
Signatures <- CELLector.createAllSignatures(CSS$navTable)</pre>
data.frame(Signatures = Signatures$S,'SubType Size'=Signatures$STS)
```

CELLector.HCCancerDrivers

High Confidence Cancer Driver genes

### Description

A list of high confidence cancer driver genes from [1]

### Usage

```
data("CELLector.HCCancerDrivers")
```

### Format

A vector of strings with one entry per cancer gene.

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#### Author(s)

Francesco Iorio (fi9232@gmail.com)

#### References

[1] Iorio, F. et al. A Landscape of Pharmacogenomic Interactions in Cancer. Cell 166, 740–754 (2016).

### **Examples**

```
data(CELLector.HCCancerDrivers)
## maybe str(CELLector.HCCancerDrivers); plot(CELLector.HCCancerDrivers) ...
```

CELLector.makeSelection

Genomics guided selection of Cancer Cell lines

### Description

Given a CELLector searching space

(outputted by the CELLector.Build\_Search\_Space function) with tumour genomic subtypes and matched underlying signatures of cancer functional events (CFEs, as defined in [1]), and a map of human cancer cell lines on it (outputted by the CELLector.buildModelMatrix function), this function selects n most representative cell lines by applying a greedy strategy described in [2] in order to maximise the covered genomic heterogeneity of primary tumours.

### Usage

CELLector.makeSelection(modelMat, n, searchSpace)

### **Arguments**

modelMat

A named binary matrix with tumour suptypes numerical identifiers on the rows, cell line names on the column and entries specifiyng whether the cell line in the column is representative of the subtype on the row (based on the collective presence/absence of the corresponding signature of CFEs). This is outputted by the CELLector.buildModelMatrix function starting from a CELLector search space (outputted by the CELLector.Build\_Search\_Space) and a cell line binary event matrix (BEM): a data frame in which the first two columns contain the COSMIC [3] identiefiers and names of cell lines (one per row), respectively, and then binary entries indicating the status of each CFEs (one per column) across cell lines. The format is the same of the entries of the list in the built-in CELLector.CellLine.BEMs object

An integer specifying the number of cell lines to select

searchSpace

A CELLector searching space, outputted by the CELLector.Build\_Search\_Space from a BEM modeling a cohort of cancer patients. With cancer functional events (CFEs) on the columns and sample identifiers on the rows. See CELLector.PrimTum.BEMs for further details

#### Value

A data frame with one row per selected cell line and the following columns:

Tumour.SubType.Index

The numerical index of the represented tumour subtype (this is the same index that the subtype has in the inputted CELLector searching space)

Representative.Cell.Line

The name of the selected cell line

Signature

The signature of CFEs underlying the subtype under consideration and collectively present in the selected cell line

percentage.patients

The size of the considered represented subtype with respect the whole cohort of cancer patients

#### Author(s)

Hanna Najgebauer and Francesco Iorio

### References

- [1] Iorio, F. et al. A Landscape of Pharmacogenomic Interactions in Cancer. Cell 166, 740–754 (2016).
- [2] Najgebauer, H. et al. Genomics Guided Selection of Cancer in vitro Models.

https://doi.org/10.1101/275032

[3] Forbes, S. A. et al. COSMIC: exploring the world's knowledge of somatic mutations in human cancer. Nucleic Acids Res. 43, D805–11 (2015).

### See Also

```
CELLector.Build_Search_Space,
CELLector.buildModelMatrix,
CELLector.CellLine.BEMs,
CELLector.PrimTum.BEMs
```

```
data(CELLector.PrimTum.BEMs)
data(CELLector.Pathway_CFEs)
data(CELLector.CFEs.CNAid_mapping)
data(CELLector.CFEs.CNAid_decode)
data(CELLector.HCCancerDrivers)
data(CELLector.CellLine.BEMs)

### Change the following two lines to work with a different cancer type
tumours_BEM<-CELLector.PrimTum.BEMs$COREAD
CELLlineData<-CELLector.CellLine.BEMs$COREAD

### unicize the sample identifiers for the tumour data
tumours_BEM<-CELLector.unicizeSamples(tumours_BEM)

### building a CELLector searching space focusing on three pathways</pre>
```

```
### and TP53 wild-type patients only
CSS<-CELLector.Build_Search_Space(ctumours = t(tumours_BEM),</pre>
                                   verbose = FALSE,
                                   minGlobSupp = 0.05,
                                   cancerType = 'COREAD',
                                   pathwayFocused = c("RAS-RAF-MEK-ERK / JNK signaling",
                                                       "PI3K-AKT-MTOR signaling",
                                                       "WNT signaling"),
                                   pathway_CFEs = CELLector.Pathway_CFEs,
                                   cnaIdMap = CELLector.CFEs.CNAid_mapping,
                                   cnaIdDecode = CELLector.CFEs.CNAid_decode,
                                   cdg = CELLector.HCCancerDrivers,
                                   subCohortDefinition='TP53',
                                   NegativeDefinition=TRUE)
### take all the signatures from the searching space
Signatures <- CELLector.createAllSignatures(CSS$navTable)</pre>
### mapping the cell lines on the CELLector searching space
ModelMat<-CELLector.buildModelMatrix(Signatures$ES,CELLlineData,CSS$navTable)
### selecting 10 cell lines
selectedCellLines<-CELLector.makeSelection(modelMat = ModelMat,</pre>
                                            n=10.
                                            searchSpace = CSS$navTable)
selectedCellLines
```

CELLector.mostSupported\_CFEs

Most recurrent combinations of Cancer Functional Events

### **Description**

This function identifies the most frequent combination of cancer functional events (CFEs) in a large cohort of cancer patients.

### Usage

### **Arguments**

transactions

A named binary matrix with CFEs on the rows, samples on the columns and entries specifying the presence/absence of a given CFE in a given sample: the *transactions* object.

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minSupport The minimal support that a combination of CFEs must have, i.e. the minimal

ratio of samples in which the CFEs must be observed simoultanously, in order

to be considered in the analysis.

minlen The minimal length of a combination of CFEs (of how many indivudal CFE it

needs to be composed) in order to be considered in the analysis (1 by default).

maxLen The maximal length of a combination of CFEs (the maximal number of indivu-

dal CFEs) in order to be considered in the analysis (10 by default).

#### **Details**

This function uses the *eclat* function from the R package *arules*.

### Value

A list with the following fields:

MSIS A string or a vector of strings (depending on the argument minlen) specifying

the CFE (or the combination of individual CFEs) that is the most frequently

observed (simultaneously across the samples in input)

SUPPORT The ratio of samples where the combination of CFEs in MSIS is obaserved on the

total number of samples, i.e. number of columns in the transactions argument

absSUPPORT The number of samples where the combination of CFEs in MSIS is obaserved

supportingSamples

The identifiers of the samples supporting MSIS, i.e. the names of the columns of *transactions*, in which the entries corresponding to MSIS rows are equal to 1.

### Author(s)

Hanna Najgebauer and Francesco Iorio

### References

Najgebauer et al., CELLector: Genomics Guided Selection of Cancer in vitro Models. doi:10.1101/275032

### **Examples**

CELLector. MSI status Cell lines' Microsatellite status

### **Description**

The microsatellite status of the cell lines in the CELLector collection, which can be stable (MSI-S), lowly instable (MSI-L), or highly instable (MSI-H) from [1]

### Usage

```
data("CELLector.MSIstatus")
```

#### **Format**

A named vector of string with one entry per cell lines (with COSMIC [2] identifiers as names) specigying the MSI status of each cell line as detailed in the description above.

### Author(s)

Francesco Iorio (fi9232@gmail.com)

#### References

- [1] Iorio, F. et al. A Landscape of Pharmacogenomic Interactions in Cancer. Cell 166, 740–754 (2016).
- [2] Forbes, S. A. et al. COSMIC: exploring the world's knowledge of somatic mutations in human cancer. Nucleic Acids Res. 43, D805–11 (2015

### **Examples**

```
data(CELLector.MSIstatus)
head(CELLector.MSIstatus)
```

CELLector.Pathway\_CFEs

Cancer functional events in biological pathways

### **Description**

Lists of cancer functional events (CFEs) from [1] involving genes in 14 key cancer biological pathways

### Usage

```
data("CELLector.Pathway_CFEs")
```

### **Format**

Named list of string vectors, whose elements are CFEs involving genes in a fixed biological pathway.

### Author(s)

Francesco Iorio (fi9232@gmail.com)

#### References

[1] Iorio, F. et al. A Landscape of Pharmacogenomic Interactions in Cancer. Cell 166, 740–754 (2016).

```
data(CELLector.Pathway_CFEs)
CELLector.Pathway_CFEs$`RAS-RAF-MEK-ERK / JNK signaling`
```

CELLector.PrimTum.BEMs

Primary Tumours' Binary Event Matrices

### **Description**

A list containing 16 data frames (one for cancer type), identified through TCGA labels. Each of these data frames contains primary tumours' *binary event matrices* (BEMs) with the status (presence/absence) of *cancer functional events* (CFEs) as defined in [1].

### Usage

data("CELLector.PrimTum.BEMs")

#### **Format**

A named list of binary matrices (with TCGA cancer type labels as names). The entries of each of these matrices indicate the status (Present/Absent) of each CFE (one per row) across primary tumors samples (one per column).

#### **Details**

BEMs of primary tumours from the Genomics of Drug Sensitivity in Cancer (GDSC1000, [1]) study. Data is available for 16 different TCGA cancer types: *BLCA, BRCA, COREAD, GBM, HNSC, KIRC, LAML, LGG, LUAD, LUSC, OV, PRAD, SKCM, STAD, THCA, UCEC*.

A decoding table for these labels is available at

https://www.cancerrxgene.org/gdsc1000/GDSC1000\_WebResources/Data\_Summary.html

Each data frame contains primary tumour samples on the columns and CFEs on the rows, with entries indicating the presence/absence of a given CFE in a given primary tumour sample.

Gene symbols as row names indicate high confidence cancer driver genes and the entries in the corresponding rows indicate the presence/absence of somatic mutations. Row names with *cna* as prefix indicate chromosomal segments that are recurrently copy number altered in cancer (RACSs, defined in [1]). A list with all the considered CFEs is available in the CELLector.CFEs data object. A decoding table for the RACSs is available in the CELLector.CFEs.CNAid\_decode, with the mapping realised by the values in the CNA\_identifier column.

Please note that the same RACS identifier across multiple cancer types might indicate different chromosomal regions, therefore in order to be decode it should be considered jointly with the TCGA label of the data frame it has been extracted from.

### Author(s)

Francesco Iorio (fi9232@gmail.com)

### References

[1] Iorio, F. et al. A Landscape of Pharmacogenomic Interactions in Cancer. Cell 166, 740–754 (2016).

### See Also

CELLector.CellLine.BEMs, CELLector.CFEs, CELLector.CFEs.CNAid\_decode

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#### **Examples**

```
data(CELLector.PrimTum.BEMs)
CELLector.PrimTum.BEMs$COREAD[c('BRAF','KRAS','cna27'),1:10]
```

CELLector.Score

Cell line quality score

### **Description**

This function computes a quality score of each cell line in terms of its ability to represent an entire cohort of considered disease matching patients. This is quantified as a trade-off between two factors. The first factor is the length of the CELLector signatures (in terms of number of composing individual alterations) that are present in the cell line under consideration. This is proportional to the granularity of the representative ability of the cell line, i.e. the longest the signature the more precisely defined is the represented sub-cohort of patients. The second factor is the size of the patient subpopulation represented by the signatures that can be observed in the cell line under consideration, thus accounting for the prevalence of the sub-cohort modeled by that cell line. An input parameter allows for these two factor to be weighted equally or differently.

### Usage

```
CELLector.Score(NavTab, CELLlineData, alpha = 0.75)
```

### **Arguments**

NavTab A CELLector searching space encoded as binary tree in a navigable table, as

returned by the CELLector.Build\_Search\_Space function.

CELLlineData A data frame in which the first two columns contain the COSMIC [1] identiefiers

(or any other alphanumeric identifier) and names of cell lines (one per row), respectively, and then binary entries indicating the status of cancer functional events (CFEs, as defined in [2], one per column) across cell lines. The format is the same of the entries of the list in the built-in CELLector.CellLine.BEMs. Based on this data the cell lines are mapped onto the CELLector searching space

and their quality assessed.

alpha A parameter that weights the contribution of two factors accounted to compute

the quality score of a cell line: respectively, the length of a signature that can be observed in a given cell line (weight = alpha) and the ratio of the sub-cohort of patients represented by that signature (1 - alpha). A cell line can be positive for multiple signatures, thus resulting in multiple quality scores. The largest is

selected for each cell line.

#### Value

A data frame with one row per cell line and the following columns:

CellLines containing an alphanumerical identifier of each cell line. These are the same as

in CELLlineData;

GlobalSupport the ratio of patients represented by the signature yielding the best CELLectorScore

for the cell line under consideration;

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SignatureLength

the length of the signature (in terms of number of individual CFEs [2]) yielding

the best  ${\tt CELLectorScore}$  for the cell line under consideration;

**CELLectorScores** 

the CELLector quality scores;

Signature The signature yielding the best CELLector quality score for the cell line under

consideration.

#### Author(s)

Francesco Iorio (fi9232@gmail.com)

#### References

[1] Forbes, S. A. et al. COSMIC: exploring the world's knowledge of somatic mutations in human cancer. Nucleic Acids Res. 43, D805–11 (2015).

[2] Iorio, F. et al. A Landscape of Pharmacogenomic Interactions in Cancer. Cell 166, 740–754 (2016).

#### See Also

CELLector.Build\_Search\_Space

```
data(CELLector.PrimTum.BEMs)
data(CELLector.Pathway_CFEs)
data(CELLector.CFEs.CNAid_mapping)
data(CELLector.CFEs.CNAid_decode)
data(CELLector.HCCancerDrivers)
data(CELLector.CellLine.BEMs)
### Change the following two lines to work with a different cancer type
tumours_BEM<-CELLector.PrimTum.BEMs$COREAD</pre>
CELLlineData<-CELLector.CellLine.BEMs$COREAD
### unicize the sample identifiers for the tumour data
tumours_BEM<-CELLector.unicizeSamples(tumours_BEM)</pre>
### building a CELLector searching space focusing on three pathways
### and TP53 wild-type patients only
CSS<-CELLector.Build_Search_Space(ctumours = t(tumours_BEM),</pre>
                                   verbose = FALSE,
                                   minGlobSupp = 0.05,
                                   cancerType = 'COREAD',
                                   pathwayFocused = c("RAS-RAF-MEK-ERK / JNK signaling",
                                                       "PI3K-AKT-MTOR signaling",
                                                       "WNT signaling"),
                                   pathway_CFEs = CELLector.Pathway_CFEs,
                                   cnaIdMap = CELLector.CFEs.CNAid_mapping,
                                   cnaIdDecode = CELLector.CFEs.CNAid_decode,
                                   cdg = CELLector.HCCancerDrivers,
                                   subCohortDefinition='TP53',
```

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### NegativeDefinition=TRUE)

```
### Evaluating cell line quality
CScores<-CELLector.Score(NavTab=CSS$navTable,CELLlineData = CELLlineData)
### Visualising best cell lines
head(CScores)</pre>
```

CELLector.selectionVisit

Visiting the CELLector searching space

### Description

This function visits the CELLector searching space using the greedy algorithm described in [1] and it returns the cancer patients subtypes identifiers sorted for cell line selection, maximising the genomic heterogeneity observed in the analysed cohort of cancer patients when considering the first n subtypes.

### Usage

```
CELLector.selectionVisit(TAV)
```

### **Arguments**

TAV

A CELLector searching space stored as a *Navigable Table*, outputted by the CELLector.Build\_Search\_Space function

### Value

A numerical vector containing the subtypes identifers sorted as explained above

### Author(s)

Hanna Najgebauer and Francesco Iorio

### References

[1] Najgebauer, H. et al. Genomics Guided Selection of Cancer in vitro Models. https://doi.org/10.1101/275032

### See Also

```
CELLector.Build_Search_Space
```

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#### **Examples**

```
data(CELLector.PrimTum.BEMs)
data(CELLector.Pathway_CFEs)
data(CELLector.CFEs.CNAid_mapping)
data(CELLector.CFEs.CNAid_decode)
data(CELLector.HCCancerDrivers)
data(CELLector.CellLine.BEMs)
tumours_BEM<-CELLector.PrimTum.BEMs$COREAD
CELLlineData<-CELLector.CellLine.BEMs$COREAD
### unicize the sample identifiers for the tumour data
tumours_BEM<-CELLector.unicizeSamples(tumours_BEM)</pre>
### building the CELLector searching space
CSS<-CELLector.Build_Search_Space(ctumours = t(tumours_BEM),</pre>
                                   verbose = FALSE,
                                   minGlobSupp = 0.05,
                                   cancerType = 'COREAD',
                                   pathway_CFEs = CELLector.Pathway_CFEs,
                                   cnaIdMap = CELLector.CFEs.CNAid_mapping,
                                   mutOnly = FALSE,
                                   cnaIdDecode = CELLector.CFEs.CNAid_decode,
                                   cdg = CELLector.HCCancerDrivers)
CELLector.selectionVisit(CSS$navTable)
```

CELLector.solveFormula

Identify cell lines harbouring a signature of Cancer Functional Events

### **Description**

This function takes in input a signature of Cancer Functional Events (CFEs, defined in [1]) as outputted by the CELLector.createAllSignatures function, and a binary event matrix (BEM) modeling the presence/absence of all the CFEs across a set of immortalised human cancer cell lines. It returns the set of cell line collectively harbouring the inputted signature.

### Usage

```
CELLector.solveFormula(RULE, dataset, To_beExcluded = NULL)
```

### Arguments

RULE A string representing a signature of cancer functional events (CFEs, defined in

[1]), i.e. names of CFEs space separated (and possibly negated ~)

dataset A data frame in which the first two columns contain the COSMIC [2] identiefiers

and names of cell lines (one per row), respectively, and then binary entries indicating the status of each CFEs (one per column) across cell lines. The format is the same of the entries of the list in the built-in CELLector.CellLine.BEMs

object.

To\_beExcluded If different from NULL (default value), then this must be a list of strings with

cell line names that should be excluded a priori from the output.

#### Value

A list with the following entries:

PS Positive samples: names of the cell lines collectively harbouring the signature

of CFEs provided in input

N The number of cell lines in PS

PERC The number of cell lines collectively harbouring the signatures of CFEs provided

in input as ratio of the total number of cell lines in the inputted BEM

#### Author(s)

Hanna Najgebauer and Francesco Iorio

#### References

[1] Iorio, F. et al. A Landscape of Pharmacogenomic Interactions in Cancer. Cell 166, 740–754 (2016).

[2] Forbes, S. A. et al. COSMIC: exploring the world's knowledge of somatic mutations in human cancer. Nucleic Acids Res. 43, D805–11 (2015).

#### See Also

```
CELLector.createAllSignatures, CELLector.CellLine.BEMs
```

### **Examples**

CELLector.unicizeSamples

Unicize patient samples' identifiers

### **Description**

This function checks if there are multiple samples derived from the same patients in the binary event matrix (BEM) modeling the presence/absence of the cancer functional events (CFEs, defined in [1]), in the cancer patients. These can be mantained (and in this case their identifier will be made unique) or discarded

### Usage

#### **Arguments**

ctumours A binary matrix with entries indicating the status (Present/Absent) of each CFE

(one per row) across primary tumors samples (one per column).

keepReplicates A boolean value indicating whether the duplicated samples should be kept (and

their identifier made unique, by adding a progressive numerical suffix) or discarded (in this case only one sample per patient will be kept and identifiers

unchanged).

### Value

A binary matrix with entries indicating the status (Present/Absent) of each CFE (one per row) across primary tumors samples (one per column), and with unique patients' (column) identifiers

### Author(s)

Hanna Najgebauer and Francesco Iorio

#### References

[1] Iorio, F. et al. A Landscape of Pharmacogenomic Interactions in Cancer. Cell 166, 740–754 (2016).

#### See Also

```
CELLector.PrimTum.BEMs
```

### **Examples**

```
data(CELLector.PrimTum.BEMs)

tumours_data<-CELLector.PrimTum.BEMs$COREAD

dim(tumours_data)
length(unique(colnames(tumours_data)))

tumours_data<-CELLector.unicizeSamples(tumours_data)

dim(tumours_data)
length(unique(colnames(tumours_data)))</pre>
```

CELLector.visualiseSearchingSpace

Visualising the CELLector searching space as a collapsibe tree

### **Description**

This function visualise the CELLector searching space as an interactive collapsible tree. Moving the mouse over the nodes of this tree shows patients' subtype details and (optionally) matched cell lines

### Usage

```
CELLector.visualiseSearchingSpace(searchSpace, CLdata = NULL)
```

### **Arguments**

searchSpace A CELLector searching space as returned by the CELLector.Build\_Search\_Space

function

CLdata If different from NULL (default) this argument must contain a data frame in

which the first two columns contain the COSMIC [1] identiefiers and names of cell lines (one per row), respectively, and then binary entries indicating the status of cancer functional events (CFEs, as defined in [2], one per column) across cell lines. The format is the same of the entries of the list in the built-in CELLector.CellLine.BEMs. Based on this data the cell lines are mapped onto the CELLector searching space and the number of cell lines mapped in each

subtype showed when moving the mouse on the corresponding node.

#### Author(s)

Hanna Najgebauer and Francesco Iorio

#### References

- [1] Forbes, S. A. et al. COSMIC: exploring the world's knowledge of somatic mutations in human cancer. Nucleic Acids Res. 43, D805–11 (2015).
- [2] Iorio, F. et al. A Landscape of Pharmacogenomic Interactions in Cancer. Cell 166, 740–754 (2016).

#### See Also

```
CELLector.Build_Search_Space, CELLector.CellLine.BEMs
```

```
data(CELLector.PrimTum.BEMs)
data(CELLector.Pathway_CFEs)
data(CELLector.CFEs.CNAid_mapping)
data(CELLector.CFEs.CNAid_decode)
data(CELLector.HCCancerDrivers)
data(CELLector.CellLine.BEMs)
tumours_BEM<-CELLector.PrimTum.BEMs$COREAD
CELLlineData<-CELLector.CellLine.BEMs$COREAD
### unicize the sample identifiers for the tumour data
tumours_BEM<-CELLector.unicizeSamples(tumours_BEM)</pre>
### building the CELLector searching space
CSS<-CELLector.Build_Search_Space(ctumours = t(tumours_BEM),</pre>
                                   verbose = FALSE,
                                   minGlobSupp = 0.05,
                                   cancerType = 'COREAD',
                                   pathway_CFEs = CELLector.Pathway_CFEs,
                                   cnaIdMap = CELLector.CFEs.CNAid_mapping,
                                   mutOnly = FALSE,
                                   cnaIdDecode = CELLector.CFEs.CNAid_decode,
                                   cdg = CELLector.HCCancerDrivers)
```

```
### visualising the Searching space as collapsible tree
CELLector.visualiseSearchingSpace(searchSpace = CSS,CLdata = CELLlineData)
```

```
CELLector.visualiseSearchingSpace_sunBurst
```

Visualising the CELLector searching space as a sunburst

### **Description**

This function visualise the CELLector searching space as an interactive sunburst.

### Usage

```
CELLector.visualiseSearchingSpace_sunBurst(searchSpace)
```

#### **Arguments**

searchSpace

A CELLector searching space as returned by the CELLector .  $\label{lector:build_Search_Space} \mbox{function}$ 

### Author(s)

Hanna Najgebauer and Francesco Iorio

### See Also

```
CELLector.Build_Search_Space,
```

```
data(CELLector.PrimTum.BEMs)
data(CELLector.Pathway_CFEs)
data(CELLector.CFEs.CNAid_mapping)
data(CELLector.CFEs.CNAid_decode)
data(CELLector.HCCancerDrivers)
data(CELLector.CellLine.BEMs)
tumours_BEM<-CELLector.PrimTum.BEMs$COREAD
CELLlineData<-CELLector.CellLine.BEMs$COREAD
### unicize the sample identifiers for the tumour data
tumours_BEM<-CELLector.unicizeSamples(tumours_BEM)</pre>
### building the CELLector searching space
CSS<-CELLector.Build_Search_Space(ctumours = t(tumours_BEM),</pre>
                                   verbose = FALSE,
                                   minGlobSupp = 0.05,
                                   cancerType = 'COREAD',
                                   pathway_CFEs = CELLector.Pathway_CFEs,
                                   cnaIdMap = CELLector.CFEs.CNAid_mapping,
                                   mutOnly = FALSE,
                                   cnaIdDecode = CELLector.CFEs.CNAid_decode,
```

cdg = CELLector.HCCancerDrivers)

### visualising the Searching space as sunburst
CELLector.visualiseSearchingSpace\_sunBurst(searchSpace = CSS)

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