# Package 'FIREVAT'

October 7, 2019

```
Type Package
Title FIREVAT, FInding REliable Variants without ArTifacts
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      which uses mutational signatures to identify sequencing artifacts and
     low-quality variants.
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Authors Andy Jinseok Lee, Hyunbin Kim
Maintainer Andy Jinseok Lee <jinseok.lee@ncc.re.kr>, Hyunbin Kim <khb7840@ncc.re.kr>
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```

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# **R** topics documented:

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Anno	tateVCFObj AnnotateVCFObj	

### **Description**

Annotates a vcf.obj using df.variants.of.interest (from PrepareAnnotationDB)

#### Usage

```
AnnotateVCFObj(vcf.obj, df.annotation.db, columns.to.include,
  include.all.columns = FALSE)
```

# Arguments

```
vcf.obj ReadVCF

df.annotation.db

A data.frame from PrepareAnnotationDB. This data.frame must have the columns 'CHROM', 'POS', 'REF', 'ALT'

columns.to.include

A character vector of columns to include. Note that existing columns in vcf.obj will not be affected.

include.all.columns
```

A boolean value. If TRUE, then annotates vcf.obj with all columns present in df.variants.of.interest. If FALSE, columns.to.include must be supplied.

#### Value

An annotated vcf.obj

CheckIfVariantRefinementIsNecessary

*CheckIfVariantRefinementIsNecessary* 

#### **Description**

Checks if variant refinement is necessary by identifying mutational signatures related to sequencing artifact in the vcf.obj (set of original unrefined point mutations).

#### Usage

```
CheckIfVariantRefinementIsNecessary(vcf.obj, bsg, df.mut.pat.ref.sigs,
  target.mut.sigs, sequencing.artifact.mut.sigs,
  init.artifact.stop = 0.05, verbose = TRUE)
```

# Arguments

vcf.obj A list from ReadVCF

bsg BSgenome.Hsapiens.UCSC object

df.mut.pat.ref.sigs

A data.frame from MutPatParseRefMutSigs

target.mut.sigs

A character vector of target mutational signatures from reference mutational signatures.

sequencing.artifact.mut.sigs

A character vector of sequencing artifact mutational signatures from reference mutational signatures.

init.artifact.stop

Numeric value less than 1. If the sum of sequencing artifact weights in vcf.obj is less than or equal to this value then this function returns judgment = FALSE, otherwise returns judgment = TRUE.

verbose

If TRUE, provides process detail. Default value is TRUE.

#### Value

A list with the following elements

- judgmentA boolean value
- seq.art.sigs.weights.sumA numeric value. Sum of sequencing artifact weights.

6 ComputeZScore

Chromosome.Names

Constant

# Description

Chromosome names for FIREVAT. Chromosome names should be given in the format of "chr" + chromosome number.

### Usage

Chromosome.Names

#### **Format**

An object of class character of length 25.

ComputeZScore

ComputeZScore

### Description

Returns a z-score of x given a distribution of values

### Usage

```
ComputeZScore(values, x)
```

# Arguments

values a numeric vector x a numeric value

### Value

a numeric value corresponding to the z-score of x

 ${\tt Compute ZScore EquiValue}$ 

ComputeZScoreEquiValue

### Description

Returns a numeric value that is equivalent to the specified z.score in the distribution of 'values'

### Usage

```
ComputeZScoreEquiValue(z.score, values)
```

### **Arguments**

z.score numeric value values numeric vector

#### Value

a numeric value corresponding to the specified z.score in the 'values' distribution

DecimalCeiling

Decimal Ceiling

### Description

Returns the ceiling of a decimal value e.g. value = 0.15, decimal = 0.1 returns 0.2

### Usage

```
DecimalCeiling(value, decimal)
```

### Arguments

value numeric value (decimal)

decimal numeric value (e.g. 0.1, 0.001)

#### Value

a numeric value

DefaultFilterToBinary

Default.Obj.Fn Default.Obj.Fn

#### **Description**

Calculates the default objective value for FIREVAT GA optimization.

### Usage

```
Default.Obj.Fn(C.refined, A.refined, C.artifactual, A.artifactual)
```

#### **Arguments**

C.refined A numeric value between 0 and 1.

A.refined A numeric value between 0 and 1.

C.artifactual A numeric value between 0 and 1.

A.artifactual A numeric value between 0 and 1.

#### Value

A numeric value between 0 and 1.

DefaultFilterToBinary Transform default filtering parameters to a binary vector

### Description

This function transforms default filtering parameter to binary vector which can be used as a suggested solution in GA algorithm.

### Usage

```
DefaultFilterToBinary(vcf.filter, params.bit.len)
```

# **Arguments**

```
vcf.filter A list generated in MakeFilter
params.bit.len A list with bit lengths of filtering parameters which is generated from ParameterToBits
```

### Value

A binary vector

Enumerate Tri Nuc Counts 9

EnumerateTriNucCounts EnumerateTriNucCounts

#### **Description**

Returns C>A, C>G, C>T, T>A, T>C, T>G counts

#### **Usage**

EnumerateTriNucCounts(spectrum)

#### **Arguments**

spectrum

a numeric vector with 96 numeric values

#### **Details**

Please note that this function assumes that 'spectrum' is sorted (i.e.  $1:16 \rightarrow C>A$ ;  $17:32 \rightarrow C>G$ ;  $33:48 \rightarrow C>T$ ;  $49:64 \rightarrow T>A$ ;  $65:80 \rightarrow T>C$ ;  $81:96 \rightarrow T>G$ )

#### Value

a numeric vector of length 6 corresponding to the counts of each trinucleotide change (C>A, C>G, C>T, T>A, T>C, T>G)

```
Euc.Exp.Weighted.Obj.Fn
```

Euc.Exp.Weighted.Obj.Fn

### Description

Calculates the Euclidean-distance of logarithmically weighted objective value for FIREVAT GA optimization.

#### Usage

```
Euc.Exp.Weighted.Obj.Fn(C.refined, A.refined, C.artifactual, A.artifactual)
```

### Arguments

C.refined A numeric value between 0 and 1.

A.refined A numeric value between 0 and 1.

C.artifactual A numeric value between 0 and 1.

A.artifactual A numeric value between 0 and 1.

#### Value

A numeric value between 0 and 1.

```
Euc.Exp.Weighted.Seq.Art.Only.Obj.Fn.1

Euc.Exp.Weighted.Seq.Art.Only.Obj.Fn.1
```

### Description

Calculates the Euclidean-distance of logarithmically weighted objective value for FIREVAT GA optimization.

### Usage

```
Euc.Exp.Weighted.Seq.Art.Only.Obj.Fn.1(C.refined, A.refined, C.artifactual,
   A.artifactual)
```

### **Arguments**

C.refined A numeric value between 0 and 1.

A.refined A numeric value between 0 and 1.

C.artifactual A numeric value between 0 and 1.

A.artifactual A numeric value between 0 and 1.

#### Value

A numeric value between 0 and 1.

```
Euc.Exp.Weighted.Seq.Art.Only.Obj.Fn.2

Euc.Exp.Weighted.Seq.Art.Only.Obj.Fn.2
```

#### **Description**

Calculates the Euclidean-distance of logarithmically weighted objective value for FIREVAT GA optimization.

# Usage

```
Euc.Exp.Weighted.Seq.Art.Only.Obj.Fn.2(C.refined, A.refined, C.artifactual,
  A.artifactual)
```

```
C.refined A numeric value between 0 and 1.

A.refined A numeric value between 0 and 1.

C.artifactual A numeric value between 0 and 1.

A.artifactual A numeric value between 0 and 1.
```

Euc.Obj.Fn

### Value

A numeric value between 0 and 1.

Euc.Obj.Fn

Euc.Obj.Fn

### **Description**

Calculates the Euclidean-distance based objective value for FIREVAT GA optimization.

### Usage

```
Euc.Obj.Fn(C.refined, A.refined, C.artifactual, A.artifactual)
```

#### **Arguments**

C.refined A numeric value between 0 and 1.

A.refined A numeric value between 0 and 1.

C.artifactual A numeric value between 0 and 1.

A.artifactual A numeric value between 0 and 1.

### Value

A numeric value between 0 and 1.

```
Exp.Weighted.A.Art.Obj.Fn

Exp.Weighted.A.Art.Obj.Fn
```

### **Description**

Exponentially weighted objective function

### Usage

```
Exp.Weighted.A.Art.Obj.Fn(C.refined, A.refined, C.artifactual,
   A.artifactual)
```

```
C.refined A numeric value between 0 and 1.

A.refined A numeric value between 0 and 1.

C.artifactual A numeric value between 0 and 1.

A.artifactual A numeric value between 0 and 1.
```

#### Value

A numeric value between 0 and 1.

```
\label{eq:continuous} {\it Exp. Weighted. A. Ref. Obj. Fn} \\ {\it Exp. Weighted. A. Ref. Obj. Fn}
```

### **Description**

Exponentially weighted objective function

# Usage

```
Exp.Weighted.A.Ref.Obj.Fn(C.refined, A.refined, C.artifactual,
   A.artifactual)
```

### **Arguments**

```
C.refined A numeric value between 0 and 1.

A.refined A numeric value between 0 and 1.

C.artifactual A numeric value between 0 and 1.

A.artifactual A numeric value between 0 and 1.
```

### Value

A numeric value between 0 and 1.

```
{\tt Exp.Weighted.Obj.Fn.1} \ \ \textit{Exp.Weighted.Obj.Fn.1}
```

### **Description**

Calculates the exponentially weighted objective value for FIREVAT GA optimization.

### Usage

```
Exp.Weighted.Obj.Fn.1(C.refined, A.refined, C.artifactual, A.artifactual)
```

```
C.refined A numeric value between 0 and 1.

A.refined A numeric value between 0 and 1.

C.artifactual A numeric value between 0 and 1.

A.artifactual A numeric value between 0 and 1.
```

### Value

A numeric value between 0 and 1.

```
Exp.Weighted.Obj.Fn.2 Exp.Weighted.Obj.Fn.2
```

#### **Description**

Calculates the exponentially weighted objective value for FIREVAT GA optimization.

#### Usage

```
Exp.Weighted.Obj.Fn.2(C.refined, A.refined, C.artifactual, A.artifactual)
```

#### **Arguments**

```
C.refined A numeric value between 0 and 1.

A.refined A numeric value between 0 and 1.

C.artifactual A numeric value between 0 and 1.

A.artifactual A numeric value between 0 and 1.
```

#### Value

A numeric value between 0 and 1.

```
Exp.Weighted.Refined.Seq.Art.Only.Obj.Fn

Exp.Weighted.Refined.Seq.Art.Only.Obj.Fn
```

### Description

Calculates the Euclidean-distance of logarithmically weighted objective value for FIREVAT GA optimization.

# Usage

```
Exp.Weighted.Refined.Seq.Art.Only.Obj.Fn(C.refined, A.refined,
   C.artifactual, A.artifactual)
```

```
C.refined A numeric value between 0 and 1.

A.refined A numeric value between 0 and 1.

C.artifactual A numeric value between 0 and 1.

A.artifactual A numeric value between 0 and 1.
```

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#### Value

A numeric value between 0 and 1.

```
FilterByStrandBiasAnalysis
```

Filter By Strand Bias Analysis

# Description

Filters refined.vcf.obj by strand bias analysis and moves these filtered variants to artifactual.vcf.obj

#### Usage

```
FilterByStrandBiasAnalysis(refined.vcf.obj, artifactual.vcf.obj, perform.fdr.correction, filter.by.strand.bias.analysis.cutoff)
```

### **Arguments**

```
refined.vcf.obj

A list of vcf data
artifactual.vcf.obj

A list of vcf data
perform.fdr.correction

A boolean value.
filter.by.strand.bias.analysis.cutoff
A numeric value.
```

#### Value

A list with filtering parameter values

- refined.vcf.obj updated refined.vcf.obj
- artifactual.vcf.obj updated artifactual.vcf.obj

FilterVCF

FilterVCF

### **Description**

Filter vcf based on the filter Filtering parameters are saved in config.obj Split vcf.obj into vcf.obj.filtered & vcf.obj.artifact based on vcf.filter

#### Usage

```
FilterVCF(vcf.obj, vcf.filter, config.obj, include.array = NULL,
  force.include = FALSE, verbose = TRUE)
```

GenerateConfigObj 15

### **Arguments**

vcf.obj A list from ReadVCF

vcf.filter A list from MakeMuTect2Filter config.obj A list from ParseConfigFile

include.array A boolean vector

force.include A boolean value. If TRUE, then uses 'include.array'

verbose If true, provides process detail

#### Value

A list with the following elements

- 1) Mutations which passed filteringvcf.obj.filtered = vcf.obj (list with data, header, genome)
- 2) Mutations which did not pass filteringvcf.obj.artifact = vcf.obj (list with data, header, genome)

GenerateConfigObj Generate config.obj by checking vcf header

#### **Description**

This function generate config.obj by checking vcf header. Users should fill in the information needed in console. In current version, only Integers & Float values can be used in config.obj for running FIREVAT.

#### Usage

```
GenerateConfigObj(vcf.obj, save.config = TRUE,
  config.path = "../temp/FIREVAT_configure.json")
```

# Arguments

vcf.obj A list from ReadVCF

save.config If true, save config.obj to config.path

config.path File path to write config.obj (json or yaml)

### Value

config.obj

GetCOSMICMutSigs

**GetCOSMICMutSigs** 

#### **Description**

Returns a data.frame of the COSMIC mutational signature reference file from the data directory

### Usage

GetCOSMICMutSigs()

### Value

a data.frame of the COSMIC reference mutational signatures

 $\label{lem:getCOSMICMutSigsEtiologiesColors} Get COSMICMutSigsNames$ 

### **Description**

Returns all COSMIC mutational signature etiologies and colors

### Usage

GetCOSMICMutSigsEtiologiesColors()

#### Value

data.frame with following columns: signature, group and color.

 ${\tt GetCOSMICMutSigsNames} \quad \textit{GetCOSMICMutSigsNames}$ 

# Description

Returns all COSMIC mutational signature names

### Usage

GetCOSMICMutSigsNames()

#### Value

a character vector

GetGASuggestedSolutions

GetGASuggestedSolutions

### **Description**

Computes suggested solutions

# Usage

```
GetGASuggestedSolutions(vcf.obj, bsg, config.obj, lower.upper.list,
   df.mut.pat.ref.sigs, target.mut.sigs, sequencing.artifact.mut.sigs,
   objective.fn, original.muts.seq.art.weights.sum, ga.preemptive.killing,
   verbose = TRUE)
```

### Arguments

vcf.obj A list from ReadVCF

bsg BSgenome.Hsapiens.UCSC object

config.obj A list from ParseConfigFile

lower.upper.list

A list from GetParameterLowerUpperVector

df.mut.pat.ref.sigs

A data.frame from MutPatParseRefMutSigs

target.mut.sigs

A character vector of the target mutational signatures from reference mutational signatures.

sequencing.artifact.mut.sigs

A character vector of the sequencing artifact mutational signatures from reference mutational signatures.

objective.fn Objective value derivation function.

original.muts.seq.art.weights.sum

A numeric value. 'seq.art.sigs.weights.sum' from CheckIfVariantRefinementIs-Necessary

ga.preemptive.killing

If TRUE, then preemptively kills populations that yield greater sequencing artifact weights sum compared to the original mutatational signatures analysis

verbose If TRUE, provides process detail. Default value is TRUE.

#### Value

A list with the following elements

- judgmentA boolean value
- seq.art.sigs.weightsA numeric value. Sum of sequencing artifact weights.

 ${\tt GetOptimizedSignatures}$ 

GetOptimizedSignatures

### **Description**

This function fetches the last row from the optimization iteration log and returns the target and artifactual mutational signatures for the type of mutations ('refined' or 'artifactual')

#### Usage

```
GetOptimizedSignatures(data, mutations.type = "refined",
    signatures = "all")
```

### **Arguments**

data A list of main data from RunFIREVAT

mutations.type A string for type of mutations ('refined' or 'artifact')

signatures A string ('all', 'target', 'artifact')

#### Value

A data.frame with the columns 'signature' and 'weight'

 ${\tt GetParameterLowerUpperVector}$ 

GetParameterLowerUpperVector

### Description

Return a lower/upper vector needed to conduct FIREVAT GA real-valued optimization.

#### Usage

```
GetParameterLowerUpperVector(vcf.obj, config.obj, vcf.filter,
  multiplier = 100)
```

vcf.obj	A list from ReadVCF
config.obj	A list from ParseConfigFile
vcf.filter	A list from MakeMuTect2Filter
multiplier	A multiplier for convert fraction to integer (default = 100)

GetPCAWGMutSigs 19

#### **Details**

vcf.obj\$data: if max(vcf.obj\$data[[param]]) < 1, then multiply multiplier to the vector

#### Value

A list with the elements

- lower.vector A numeric vector. Each element is the minimum value of each parameter
- upper.vector A numeric vector. Each element is the maximum value of each parameter
- · vcf.obj vcf.obj with updated data

GetPCAWGMutSigs

**GetPCAWGMutSigs** 

#### **Description**

Returns the PCAWG mutational signatures data

### Usage

```
GetPCAWGMutSigs(sequencing.type = "wes")
```

### **Arguments**

sequencing.type

A string value. It can be either 'wes' for whole-exome sequencing or 'wgs' for whole-genome sequencing

### Value

a data.frame of the PCAWG mutatioanl signatures

```
{\tt GetPCAWGMutSigsEtiologiesColors}
```

GetPCAWGMutSigsEtiologiesColors

### **Description**

Returns the PCAWG mutational signatures etiologies and colors

# Usage

```
GetPCAWGMutSigsEtiologiesColors()
```

#### Value

```
a data.frame with the columns 'signature', 'group', 'color'
```

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GetPCAWGMutSigsNames GetPCAWGMutSigsNames

### Description

Returns the PCAWG mutational signatures names

### Usage

```
GetPCAWGMutSigsNames()
```

#### Value

a character vector of the PCAWG mutational signatures names

InitializeVCF

*InitializeVCF* 

### Description

Initialize VCF with FIREVAT config file This functions selects point mutations and appends filter values to vcf.obj\$data

### Usage

```
InitializeVCF(vcf.obj, config.obj, verbose = TRUE)
```

### Arguments

vcf.obj A list from ReadVCF

config.obj A list from ParseConfigFile verbose If true, provides process detail

### Value

A list with the following elements

- vcf.obj.filteredvcf.obj (high-quality vcf)
- vcf.obj.artifactvcf.obj (low-quality vcf)

```
Leaky.ReLU.A.Art.Obj.Fn
```

Leaky.ReLU.A.Art.Obj.Fn

### **Description**

Leaky ReLU objective function

### Usage

```
Leaky.ReLU.A.Art.Obj.Fn(C.refined, A.refined, C.artifactual, A.artifactual)
```

#### **Arguments**

C.refined A numeric value between 0 and 1.

A.refined A numeric value between 0 and 1.

C.artifactual A numeric value between 0 and 1.

A.artifactual A numeric value between 0 and 1.

#### Value

A numeric value between 0 and 1.

```
Leaky.ReLU.A.Ref.Obj.Fn
```

Leaky.ReLU.A.Ref.Obj.Fn

# Description

Leaky ReLU objective function

#### Usage

```
Leaky.ReLU.A.Ref.Obj.Fn(C.refined, A.refined, C.artifactual, A.artifactual)
```

# Arguments

C.refined A numeric value between 0 and 1.

A.refined A numeric value between 0 and 1.

C.artifactual A numeric value between 0 and 1.

A.artifactual A numeric value between 0 and 1.

#### Value

A numeric value between 0 and 1.

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Leaky.ReLU.Obj.Fn

Leaky.ReLU.Obj.Fn

### Description

Lkeay ReLU objective function

### Usage

```
Leaky.ReLU.Obj.Fn(C.refined, A.refined, C.artifactual, A.artifactual)
```

### Arguments

C.refined A numeric value between 0 and 1.

A.refined A numeric value between 0 and 1.

C.artifactual A numeric value between 0 and 1.

A.artifactual A numeric value between 0 and 1.

#### Value

A numeric value between 0 and 1.

MakeFilter

MakeFilter

### **Description**

Creates a vcf filter from config.obj

### Usage

```
MakeFilter(config.obj)
```

# **Arguments**

config.obj

A list from ParseConfigFile (any filter with "use\_in\_filter" value declared as FALSE is not considered)

#### Value

A list with the filter parameters

MutaliskParseVCFObj

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MutaliskParseVCFObj MutaliskParseVCFObj

### **Description**

Parses a vcf.obj and prepares it to run Mutalisk.

#### Usage

```
MutaliskParseVCFObj(vcf.obj)
```

### **Arguments**

vcf.obj

A list from ReadVCF

#### Value

A data.frame

MutPatParseRefMutSigs MutPatParseRefMutSigs

#### **Description**

Parses a df.ref.mut.sigs and prepares it to run Mutational Patterns.

#### Usage

```
MutPatParseRefMutSigs(df.ref.mut.sigs, target.mut.sigs,
    signature.start.column.index = 4,
    mutation.type.header = "SomaticMutationType")
```

# Arguments

```
df.ref.mut.sigs

A data.frame of reference mutational signatures

target.mut.sigs

A character vector of target mutational signatures names

signature.start.column.index

= An integer value (e.g. column index corresponding to 'SBS1')

mutation.type.header

= A string value (name of header corresponding to column containing 'A[C>A]A'
data))
```

#### Value

A data.frame of the format deconstructSigs::signatures.cosmic

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tPatParseVCFObj MutPatParseVCFObj

# Description

Parses a vcf.obj and prepares it to run Mutational Patterns.

### Usage

```
MutPatParseVCFObj(vcf.obj, bsg, sample.id = "sample")
```

#### **Arguments**

vcf.obj A list from ReadVCF

bsg BSgenome.Hsapiens.UCSC.hg19::BSgenome.Hsapiens.UCSC.hg19 or BSgenome.Hsapiens.UCSC.hg38

sample.id A string value

#### Value

A data.frame with the column sample.id and row names corresponding to 96 substitution types

# Description

Calculate the number of bits needed to conduct FIREVAT GA binary optimization.

### Usage

```
ParameterToBits(vcf.obj, config.obj, vcf.filter, multiplier = 100)
```

# Arguments

vcf.obj	A list from ReadVCF
config.obj	A list from ParseConfigFile
vcf.filter	A list from MakeMuTect2Filter
multiplier	A multiplier for convert fraction to integer (default = 100)

#### **Details**

vcf.obj\$data: if max(vcf.obj\$data[[param]]) < 1, then multiply multiplier to the vector

ParseConfigFile 25

### Value

A list with the elements

• params.bit.lenA numeric vector. Each element is the bit length of each parameter value

• vcf.objA vcf.obj (ReadVCF) with updated data

 ${\tt ParseConfigFile}$ 

ParseConfigFile

# Description

This function returns config.obj from JSON or YAML config file. - Check if the config file is in JSON format or YAML format - Return config.obj

### Usage

```
ParseConfigFile(config.path, verbose = TRUE)
```

### Arguments

config.path A string for config file path
verbose If true, provides process detail

#### Value

```
config.obj: list of parameters
```

### **Examples**

```
## Not run:
ParseConfigFile("example.variant.caller.json")
ParseConfigFile("example.variant.caller.json", verbose=False)
## End(Not run)
```

PCAWG.All.Sequencing.Artifact.Signatures

\*\*Constant\*\*

### Description

PCAWG mutational signatures reported to be associated with sequencing artifacts

### Usage

PCAWG.All.Sequencing.Artifact.Signatures

### **Format**

An object of class character of length 18.

PCAWG.Known.Sequencing.Artifact.Signatures

\*\*Constant\*\*

### **Description**

PCAWG mutational signatures reported to be associated with sequencing artifacts

### Usage

PCAWG.Known.Sequencing.Artifact.Signatures

### **Format**

An object of class character of length 1.

 ${\it PCAWG.Possible.Sequencing.Artifact.Signatures} \\ {\it Constant}$ 

# Description

PCAWG mutational signatures reported to be associated with sequencing artifacts

### Usage

PCAWG.Possible.Sequencing.Artifact.Signatures

### **Format**

An object of class character of length 17.

```
{\it PCAWG.Target.Mutational.Signatures} \\ {\it Constant}
```

### **Description**

PCAWG target mutational signatures reported to be unrelated to sequencing artifacts

### Usage

```
PCAWG.Target.Mutational.Signatures
```

#### **Format**

An object of class character of length 47.

PerformStrandBiasAnalysis

PerformStrandBiasAnalysis

### **Description**

Performs strand bias analysis

# Usage

```
PerformStrandBiasAnalysis(vcf.obj, ref.forward.strand.var,
  ref.reverse.strand.var, alt.forward.strand.var, alt.reverse.strand.var,
  perform.fdr.correction = TRUE, fdr.correction.method = "BH")
```

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#### Value

An updated vcf.obj

PlotMutaliskResults PlotMutaliskResults

#### **Description**

Plots Mutalisk results

### Usage

```
PlotMutaliskResults(mutalisk.results, signatures, trinuc.max.y,
  trinuc.min.y, mut.type.max.y, title, font.size.small = 8,
  font.size.med = 14)
```

### **Arguments**

```
Mutalisk.results

A list obtained from RunMutalisk
signatures
A character vector of mutational signatures names
trinuc.max.y
A numeric value (maximum y-axis value)
trinuc.min.y
A numeric value (minimum y-axis value)
mut.type.max.y
A numeric value
title
A string value
font.size.small
A numeric value
font.size.med
A numeric value
```

### Value

A ggplot object

# Examples

PlotMutationTypes 29

PlotMutationTypes

*PlotMutationTypes* 

### **Description**

Plots a horizontal barplot of mutation types

#### Usage

```
PlotMutationTypes(mutation.types = c("C>A", "C>G", "C>T", "T>A", "T>C",
   "T>G"), mutation.types.values, mutation.types.colors, max.y.val, title,
   convert.to.percentage = T, show.legend = T, font.size.small = 8,
   font.size.med = 14, plot.margin = unit(c(0.5, 0.5, 0.5, 0.5), "cm"))
```

### **Arguments**

```
mutation.types \;\; Mutation \; types; \; Default = c("C>A", "C>G", "C>T", "T>A", "T>C", "T>G")
mutation.types.values
                  Mutation count for each mutation type
mutation.types.colors
                  A color vector for indicating mutation types
max.y.val
                  y axis maximum value
                  Plot title
title
convert.to.percentage
                  if True convert y values to percentage (x 100); Default = T
                  If True, show legend; Default = T
show.legend
font.size.small
                  Small font size; Default = 8
font.size.med
                  Medium font size; Default = 14
                  Margin vector for drawing plot; Default = unit(c(0.5, 0.5, 0.5, 0.5), "cm"))
plot.margin
```

#### Value

A ggplot object

### **Examples**

```
plot.margin = unit(c(0.5, 0.5, 0.5, 0.5), "cm"))
print(p)
## End(Not run)
```

PlotOptimizationIterations

**PlotOptimizationIterations** 

#### **Description**

Plots multiple scatter plots into one figure

#### Usage

```
PlotOptimizationIterations(df, columns.to.plot, x.axis.var, x.axis.title,
  x.min, x.max, save.file, title, y.axis.title = "", y.max = 1,
  point.size = 1, connect.dots = T, plot.legend = T,
  legend.ncol = 1, font.size.med = 14, font.size.large = 16,
  plot.margin = unit(c(0.5, 0.5, 0.5, 0.5), "cm"))
```

### **Arguments**

```
df
                  A data.frame (from reading "FIREVAT_Optimization_Logs.tsv")
columns.to.plot
                  A character vector (of column names to plot)
x.axis.var
                  x axis variable
x.axis.title
                  x axis title
                  x axis maximum value
x.max
save.file
                  Filename (including full path) to which the plot will be saved
title
                  Plot title
                  y axis title; Default = ""
y.axis.title
                  y axis maximum value; Default = 1
y.max
                  Point size: Default = 1
point.size
connect.dots
                  If True draws dots for each iteration; Default = True
plot.legend
                  If True write legend of plot; Default = T
legend.ncol
                  legend.n Default = 1
font.size.med
                  Medium font size; Default = 14
font.size.large
                  Large font size; Default = 16
                  Margin vector for plot; Default = unit(c(0.5, 0.5, 0.5, 0.5), "cm"))
plot.margin
```

#### Value

A ggplot object

```
PlotSignaturesContProbs
```

*PlotSignaturesContProbs* 

# Description

Plots a horizontal barplot of identified mutational signatures

#### Usage

```
PlotSignaturesContProbs(df.identified.mut.sigs, df.ref.sigs.groups.colors,
  title, convert.to.percentage = T, font.size.small = 8,
  font.size.med = 14, plot.margin = unit(c(0.5, 0.5, 0.5, 0.5), "cm"))
```

#### **Arguments**

```
df.identified.mut.sigs
A data.frame of identified mutational signatures

df.ref.sigs.groups.colors
A data.frame with 'signature', 'group', and 'color' columns

title
Plot title
convert.to.percentage
If true, convert y values to percentage (x 100); Default = T,

font.size.small
Small font size; Default = 8,

font.size.med
Medium font size; Default = 14,

plot.margin
Margin vector for drawing plot; Default = unit(c(0.5, 0.5, 0.5, 0.5), "cm"))
```

#### Value

A ggplot object

### **Examples**

```
## Not run:
    g <- PlotSignaturesContProbs(sigs = c(mutalisk.results$identified.mut.sigs),
    sigs.probs = c(mutalisk.results$identified.mut.sigs.probs),
    df.ref.sigs.groups.colors = GetPCAWGMutSigsEtiologiesColors())
    print(g)
## End(Not run)</pre>
```

PlotTriNucSpectrum

PlotTable

**PlotTable** 

### **Description**

Plots basic statistics table

### Usage

```
PlotTable(df, padding = 20, font.size = 14)
```

### **Arguments**

df = A data, frame where the first column is header and the second column is data

value

padding size; Default = 20

font.size Font size; Default = 14

#### Value

A plot

PlotTriNucSpectrum

PlotTriNucSpectrum

### Description

Plots the spectrum of 96 trinucleotide distribution (C>A, C>G, C>T, T>A, T>C, T>G) Please note that this function assumes that both sub.types and spectrum are sorted in the following order: C>A, C>G, C>T, T>A, T>C, T>G

### Usage

```
PlotTriNucSpectrum(sub.types, spectrum, max.y.val, min.y.val, y.axis.title,
  draw.top.strip = T, draw.x.axis.labels = T, draw.y.axis.labels = T,
  draw.y.axis.title = T, font.size.small = 8, font.size.med = 14,
  plot.margin.top = 0.5, plot.margin.bottom = 0.5,
  plot.margin.left = 0.5, plot.margin.right = 0.5, title)
```

PlotVCFStatsBoxPlots 33

#### **Arguments**

```
A character vector (types of 96 trinucleotide substitutions)
sub.types
                  A numeric vector (96 elements)
spectrum
                  y axis maximum value
max.y.val
min.y.val
                 y axis minimum value
y.axis.title
                  y axis title
draw.top.strip If True then draws top strip; Default = T
draw.x.axis.labels
                  If True then draws x axis labels; Default = T
draw.y.axis.labels
                  If True then draws y axis labels; Default = T
draw.y.axis.title
                  If True then draws y axis title; Default = T
font.size.small
                  Small font size; Default = 8
font.size.med
                 Medium font size; Default = 14
plot.margin.top
                  Top margin; Default = 0.5
plot.margin.bottom
                  Bottom margin; Default = 0.5
plot.margin.left
                 Left margin; Default = 0.5
plot.margin.right
                 Right margin; Default = 0.5
title
                  Plot title
```

#### Value

A ggplot object

PlotVCFStatsBoxPlots PlotVCFStatsBoxPlots

### **Description**

Plots multiple (original, refined, artifact vcf) boxplots for single filter parameter

### Usage

```
PlotVCFStatsBoxPlots(original.vcf.stat.values, refined.vcf.stat.values,
  artifact.vcf.stat.values, xlab, axis.font.size = 10,
  label.font.size = 10, title.font.size = 12)
```

#### **Arguments**

```
original.vcf.stat.values

A numeric vector corresponding to the original vcf.obj values of single filter parameter

refined.vcf.stat.values

A numeric vector corresponding to the refined vcf.obj values of single filter parameter

artifact.vcf.stat.values

A numeric vector corresponding to the artifact vcf.obj values of single filter parameter

xlab

A string value (x-axis label)

axis.font.size

An integer value (axis font size)

title.font.size
```

#### Value

A ggboxplot

PlotVCFStatsHistograms

*PlotVCFStatsHistograms* 

An integer value (title font size)

#### **Description**

Plots multiple VCF stats histograms into one figure

### Usage

```
PlotVCFStatsHistograms(plot.values, x.axis.labels, stat.y.max.vals,
    stat.x.max.vals, sample.id, save.file, title, cutoff.values,
    plot.boxplot = F, plot.cutoff.line.color = "#D4012E",
    plot.cutoff.value.lines = F, bin.width = 1, ncol = 4, nrow = 3,
    font.size.med = 10, font.size.large = 12, plot.margin = unit(c(0.5,
    0.5, 0.5, 0.5), "cm"))
```

#### **Arguments**

```
plot.values A list of multiple numeric vectors x.axis.labels A character vector of x axis labels stat.y.max.vals
```

A numeric vector of max y-axis values

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stat.x.max.vals

A numeric vector of max x-axis values

sample.id A string value of sample ID

save.file A string value of file to which the resulting plot will be saved

title A string value of plot title

cutoff.values A numeric vector of cutoff values plot.boxplot A boolean value (default = False)

plot.cutoff.line.color

A hex string value (default = "#D4012E")

plot.cutoff.value.lines

A boolean value (default = False)

bin.width An integer value (default = 1; histogram bin width)
ncol An integer value (default = 4; ggarrange ncol)
nrow An integer value (default = 3; ggarrange nrow)

font.size.med An integer value (default = 10)

font.size.large

An integer value (default = 12)

plot.margin A list (default = unit(c(0.5, 0.5, 0.5, 0.5, 0.5), "cm"))

#### Value

A list with the following elements

- f = A ggarrange object
- graphs = A list of length 3; each element is a ggplot histogram

PrepareAnnotationDB

### Description

Prepares df.genes.of.interest from a vcf.obj (ReadVCF) of COSMIC or ClinVar vcf for AnnotateVCF0bj

#### **Usage**

PrepareAnnotationDB(annotation.vcf.obj)

### Arguments

```
annotation.vcf.obj
```

vcf.obj of COSMIC or ClinVar vcf file

#### Value

A data.frame of annotation.vcf.obj

PrepareArtifactAnnotationTable

Prepare Artifact Annotation Table

### Description

Prepares artifactual mutations annotation (filtered, queried) table

### Usage

PrepareArtifactAnnotationTable(data)

### Arguments

data

A list of elements returned from RunFIREVAT

#### Value

A data.frame

 ${\tt Prepare Artifact Strand Bias Table}$ 

Prepare Artifact Strand Bias Table

### **Description**

Prepares artifactual mutations strand biased variants table

### Usage

PrepareArtifactStrandBiasTable(data)

# Arguments

data

A list of elements returned from RunFIREVAT

### Value

A data.frame

 ${\tt Prepare Artifactual Muts Optimization Iterations Plot}$ 

 $\label{lem:prepareArtifactualMutsOptimizationIterationsPlot} PrepareArtifactual MutsOptimization Iterations Plot$ 

## Description

Prepares artifactual mutations optimization iterations plot

## Usage

 $\label{lem:prepareArtifactualMutsOptimizationIterationsPlot(data)} PrepareArtifactualMutsOptimizationIterationsPlot(data)$ 

## Arguments

data

A list of elements returned from RunFIREVAT

#### Value

A ggplot object

 ${\tt Prepare Filter Cutoffs Table}$ 

Prepare Filter Cutoffs Table

#### **Description**

Prepares filter cutoffs table for reporting

## Usage

PrepareFilterCutoffsTable(data)

#### **Arguments**

data

A list of elements returned from RunFIREVAT

#### Value

 ${\tt Prepare Genetic Algorithm Parameters Table}$ 

Prepare Genetic Algorithm Parameters Table

## Description

Prepares Genetic Algorithm parameters table

## Usage

 ${\tt Prepare Genetic Algorithm Parameters Table (data)}$ 

#### **Arguments**

data

A list of elements returned from RunFIREVAT

#### Value

A data.frame

 ${\tt PrepareIdentifiedSignaturesPlot}$ 

Prepare Identified Signatures Plot

## Description

Prepares identified signatures plot for reporting

## Usage

PrepareIdentifiedSignaturesPlot(data)

## Arguments

data

A list of elements returned from RunFIREVAT

#### Value

A ggarrange object

 ${\tt Prepare MLERe constructed Spectrum sPlot}$ 

Prepare MLE Reconstructed Spectrums Plot

## Description

Prepares MLE reconstructed spectrums plot

## Usage

PrepareMLEReconstructedSpectrumsPlot(data)

#### Arguments

data

A list of elements returned from RunFIREVAT

#### Value

A ggarrange object

 $\label{lem:prepareNucleotideSubstitutionTypesPlot} PrepareNucleotideSubstitutionTypesPlot$ 

## Description

Prepares nucleotide substitution types plot

## Usage

PrepareNucleotideSubstitutionTypesPlot(data)

#### **Arguments**

data

A list of elements returned from RunFIREVAT

#### Value

A ggarrange object

PrepareObservedSpectrumsPlot

 ${\it Prepare Observed Spectrums Plot}$ 

## Description

Prepares observed spectrums plot

## Usage

PrepareObservedSpectrumsPlot(data)

## Arguments

data

A list of elements returned from RunFIREVAT

#### Value

A ggarrange object

 ${\tt PrepareOptimizationResultsTable}$ 

Prepare Optimization Results Table

## Description

Prepares optimization results table

## Usage

PrepareOptimizationResultsTable(data)

#### **Arguments**

data

A list of elements returned from RunFIREVAT

#### Value

PrepareOptimizedVCFStatisticsPlot

Prepare Optimized VCF Statistics Plot

## Description

Prepares optimized VCF statistics plot

#### Usage

PrepareOptimizedVCFStatisticsPlot(data)

#### Arguments

data

A list of elements returned from RunFIREVAT

#### Value

A ggarrange object

 ${\tt Prepare Refined Annotation Table}$ 

Prepare Refined Annotation Table

## Description

Prepares refined mutations annotation (filtered, queried) table

## Usage

PrepareRefinedAnnotationTable(data)

## Arguments

data

A list of elements returned from RunFIREVAT

#### Value

 $\label{lem:prepareRefinedMutsOptimizationIterationsPlot} Prepare Refined MutsOptimization Iterations Plot$ 

## Description

Prepares refined mutations optimization iterations plot

## Usage

PrepareRefinedMutsOptimizationIterationsPlot(data)

## Arguments

data

A list of elements returned from RunFIREVAT

#### Value

A ggplot object

 ${\tt Prepare Refined Strand Bias Table}$ 

Prepare Refined Strand Bias Table

#### **Description**

Prepares refined mutations strand biased variants table

## Usage

PrepareRefinedStrandBiasTable(data)

## Arguments

data

A list of elements returned from RunFIREVAT

#### Value

 ${\tt PrepareResidualSpectrumsPlot}$ 

Prepare Residual Spectrums Plot

## Description

Prepares residual spectrums plot

## Usage

PrepareResidualSpectrumsPlot(data)

## Arguments

data

A list of elements returned from RunFIREVAT

#### Value

A ggarrange object

 ${\tt PrepareTrinucleotideSpectrumsTable}$ 

Prepare Trinucle ot ide Spectrums Table

## Description

Prepares trinucleotide spectrums table

## Usage

PrepareTrinucleotideSpectrumsTable(data)

#### **Arguments**

data

A list of elements returned from RunFIREVAT

#### Value

QueryAnnotatedVCF

PrintLog

PrintLog

## Description

Prints log message

#### Usage

```
PrintLog(msg, type = "INFO")
```

#### Arguments

msg String value message to print along with log type and date

type String value that represents type of this message. 'INFO' by default.

 ${\tt QueryAnnotatedVCF}$ 

Filter Annotated VCF

## Description

Annotates a vcf.obj using df.variants.of.interest (from (PrepareAnnotationDB)

#### Usage

```
QueryAnnotatedVCF(vcf.obj.annotated, filter.key.value.pairs,
  filter.condition = "AND")
```

#### **Arguments**

```
vcf.obj.annotated
```

AnnotateVCF0bj

filter.key.value.pairs

A list with the key as the column name and value as the filtering values. E.g. list("CLNSIG" = c("Pathogenic", "Pathogenic/Likely\_pathogenic"))

filter.condition

'AND' or 'OR'.

#### Value

A vcf.obj

ReadOptimizationIterationReport

ReadOptimizationIterationReport

#### **Description**

Read optimization iteration report

#### Usage

ReadOptimizationIterationReport(data)

#### Arguments

data

A list of elements returned from RunFIREVAT

#### Value

A data.frame of FIREVAT optimization logs

ReadVCF

ReadVCF

#### **Description**

Reads a .vcf file

#### Usage

```
ReadVCF(vcf.file, genome = "hg19", split.info = FALSE,
   check.chromosome.name = TRUE)
```

#### **Arguments**

vcf.file (full path of a .vcf file) genome ('hg19' or 'hg38')

 $split.in fo \qquad \ \ A \ boolean \ value. \ If \ TRUE, then \ makes \ the \ INFO \ column \ in \ the \ vcf \ as \ a \ separate$ 

column. Default value is FALSE.

check.chromosome.name

A boolean value. If TRUE, then check whether converts 'MT' to 'M' and adds 'chr' to the CHROM column. Default value is TRUE.

#### Value

A list with elements 'data', 'header', 'genome'

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 ${\tt ReportFIREVATResults} \quad \textit{ReportFIREVATResults}$ 

#### Description

Reports FIREVAT results in html format (generated from Rmd)

#### Usage

```
ReportFIREVATResults(data)
```

#### **Arguments**

data

A list of main data from RunFIREVAT

#### Value

An updated data list

RunFIREVAT

RunFIREVAT

#### **Description**

Runs FIREVAT using configuration data. Filters point mutations in the user-specified vcf file based on mutational signature identification and outputs the refined and artifact vcf files as well as metadata related to the refinement process.

#### Usage

```
RunFIREVAT(vcf.file, vcf.file.genome = "hg19", config.file,
 df.ref.mut.sigs = GetPCAWGMutSigs(),
  target.mut.sigs = GetPCAWGMutSigsNames(),
  sequencing.artifact.mut.sigs = PCAWG.All.Sequencing.Artifact.Signatures,
  num.cores = 2, output.dir, mode = "ga", init.artifact.stop = 0.05,
 objective.fn = Default.Obj.Fn, use.suggested.soln = TRUE,
  ga.type = "real-valued", ga.pop.size = 100, ga.max.iter = 100,
  ga.run = 50, ga.pmutation = 0.1, ga.preemptive.killing = FALSE,
 mutalisk = TRUE, mutalisk.method = "all",
 mutalisk.must.include.sigs = NULL,
 mutalisk.random.sampling.count = 20,
 mutalisk.random.sampling.max.iter = 10,
 perform.strand.bias.analysis = FALSE,
 filter.by.strand.bias.analysis = TRUE,
  filter.by.strand.bias.analysis.cutoff = 0.25,
  strand.bias.perform.fdr.correction = TRUE,
```

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```
strand.bias.fdr.correction.method = "BH",
ref.forward.strand.var = NULL, ref.reverse.strand.var = NULL,
alt.forward.strand.var = NULL, alt.reverse.strand.var = NULL,
annotate = FALSE, df.annotation.db = NULL,
annotated.columns.to.display = NULL,
annotation.filter.key.value.pairs = NULL,
annotation.filter.condition = "AND", write.vcf = TRUE,
report = TRUE, save.rdata = TRUE, save.tsv = TRUE,
report.format = "html", verbose = TRUE)
```

#### **Arguments**

vcf.file String value corresponding to input .vcf file. Please provide the full path.

vcf.file.genome

Genome assembly of the input .vcf file. The value should be eitehr 'hg19' or

'hg38'.

config.file String value corresponding to input configuration file. For more details please

refer to ...

df.ref.mut.sigs

A data.frame of the reference mutational signatures

target.mut.sigs

A character vector of the target mutational signatures from reference mutational signatures.

sequencing.artifact.mut.sigs

A character vector of the sequencing artifact mutational signatures from reference mutational signatures.

num. cores Number of cores to allocate

output.dir String value of the desired output directory

mode String value. The value should be either 'ga' or 'manual'.

init.artifact.stop

Numeric value less than 1. If the sum of sequencing artifact weights in the user-specified original VCF file (i.e. vcf.file) is less than or equal to this value then FIREVAT does not perform variant refinement. Default value is 0.05. Note that this option does not apply if 'mode' is 'manual'.

objective.fn Objective value derivation function. Default: Default.Obj.Fn. use.suggested.soln

Boolean value. If TRUE, then FIREVAT passes the default values of filter variables declared as 'use\_in\_filter' in the config file to the 'suggestions' parameter of the Genetic Algorithm package. If FALSE, then FIREVAT supplies NULL to the GA package 'suggestions' parameter. FIREVAT also computes baseline performance of each filter variable and uses fittest population from each variable as a suggested solution.

ga. type String value. The value should be either 'binray' or 'real-valued'.

ga.pop.size Integer value of the Genetic Algorithm 'population size' parameter. Default: 100. This value should be set based on the number of filter parameters. Recommendation: 40 per filter parameter.

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ga.max.iter Integer value of the Genetic Algorithm 'maximum iterations' parameter. Dde-

fault: 100. This value should be set based on the number of filter parameters.

Recommendation: same as 'ga.pop.size'.

ga.run Integer value of the Genetic Algorithm 'run' parameter. Default: 50. This value

should be set based on the 'ga.max.iter' parameter. Recommendation: 25 per-

cent of 'ga.max.iter'.

ga.pmutation Float value of the Genetic Algorithm 'mutation probability' parameter. Default:

0.1.

ga.preemptive.killing

If TRUE, then preemptively kills populations that yield greater sequencing artifact weights sum compared to the original mutatational signatures analysis

mutalisk If TRUE, confirm mutational signature analysis with Mutalisk. Default: TRUE.

mutalisk.method

Mutalisk signature identification method. Default: 'random.sampling'. The value can be either 'all' or 'random.sampling'. 'all' uses all target.mut.sigs to identify mutational signatures. 'random.sampling' randomly samples from target.mut.sigs to identify mutational signatures.

mutalisk.must.include.sigs

Signatures that must be included in the Mutalisk signature identification A character vector corresponding to the signature names.

mutalisk.random.sampling.count

Mutalisk random sampling count. Default: 20. The number of signatures to sample from target.mut.sigs

mutalisk.random.sampling.max.iter

Mutalisk random sampling maximum iteration. Default: 10. The number of times Mutalisk randomly samples from target.mut.sigs before determining the candidate signatures.

perform.strand.bias.analysis

If TRUE, then performs strand bias analysis.

filter.by.strand.bias.analysis

If TRUE, then filters out variants in refined vcf based on strand bias analysis results

filter.by.strand.bias.analysis.cutoff

The p.value or q value cutoff for filtering out variants.

strand.bias.perform.fdr.correction

If TRUE, then performs false discovery rate correction for strand bias analysis.

strand.bias.fdr.correction.method

A string value. Default value is 'BH'. Refer to 'p.adjust()' function method.

ref.forward.strand.var

A string value.

ref.reverse.strand.var

A string value,

alt.forward.strand.var

A string value,

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alt.reverse.strand.var

A string value,

annotate A boolean value. Default value is TRUE.

df.annotation.db

A data.frame. Please refer to PrepareAnnotationDB

annotated.columns.to.display

A character vector.

annotation.filter.key.value.pairs

A list.

annotation.filter.condition

'AND' or 'OR'.

write.vcf If TRUE, write original/refined/artifact vcfs. Default: TRUE.

report If TRUE, generate report. Default: TRUE.

save.rdata If TRUE, save rdata. Default: TRUE. save.tsv If TRUE, save tsv. Default: TRUE.

report. format The format of FIREVAT report. We currently only support 'html'.

verbose If TRUE, provides process detail. Default: TRUE.

#### Value

A list with the following elements

- f = A ggarrange object
- graphs = A list of length 3; each element is a ggplot histogram

RunGAMode RunGAMode

## Description

Runs FIREVAT ga mode

## Usage

RunGAMode(data)

#### Arguments

data A list from RunFIREVAT

#### Value

A list

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RunManualMode

RunManualMode

#### **Description**

Runs FIREVAT manual mode

#### Usage

RunManualMode(data)

#### **Arguments**

data

A list from RunFIREVAT

#### Value

A list

RunMutalisk

RunMutalisk

#### **Description**

Identifies mutational signatures using Mutalisk

#### Usage

```
RunMutalisk(vcf.obj, df.ref.mut.sigs, target.mut.sigs,
  random.sampling.candidate.mut.sigs = c(), method = "random.sampling",
  n.sample = 20, n.iter = 10, verbose = TRUE)
```

#### Arguments

```
vcf.obj A list (from firevat_vcf::ReadVCF)
```

df.ref.mut.sigs

A data.frame of reference mutational signatures

target.mut.sigs

A character vector of target mutational signatures names to identify from

random.sampling.candidate.mut.sigs

A character vector of mutational signatures names that gets appended to the list of candidate mutational signatures so that these are always considered.

method

A string value (must be either 'random.sampling' or 'all'). The method 'random.sampling' samples (without replacement) 'n.sample' number of signatures 'n.iter' number of times and runs the candidate signatures one last time. The method 'all' uses all target.mut.sigs

RunMutaliskHelper 51

n.sample	An integer value ('random.sampling' method parameter) Number of signatures to choose for each iteration of random sampling).
n.iter	An integer value ('random.sampling' method parameter). Number of iterations to perform random sampling.
verbose	If true, provides process details

#### Value

A list with the following elements

- num.point.mutationsAn integer value count of total point mutations
- sub.typesA character vector of length 96
- sub.types.spectrumA numeric vector of length 96
- num.mut.sigsAn integer value (count of unique mutational signatures identified)
- · identified.mut.sigsA character vector where each element is a mutational signature identified
- identified.mut.sigs.probsA numeric vector where each element is the weight of mutational signature identified. The ordering follows identified.mut.sigs
- identified.mut.sigs.spectrumA numeric vector of length 96
- residualsA numeric vector of length 96
- rssA numeric value (residual sum of squares)
- cos.sim.scoreA numeric value (cosine similarity score between observed mutational spectrum and reconstructed mutational signatures)
- all.models.sigsA list where each element is a model; a model is a list of signatures identified)
- all.models.sigs.probsA list where each element is a model; a model is a list of contribution probabilities
- all.models.cos.sim.scoresA list where each element is a model; a model is a list of cosine similarity socres

lper RunMutaliskHelper	
ipo.	

## Description

Helper function for RunMutalisk

#### Usage

RunMutaliskHelper(vcf.trinucleotide.data, df.ref.mut.sigs, target.mut.sigs)

52 RunMutPat

#### **Arguments**

```
vcf.trinucleotide.data
A data.frame (from firevat_mutalisk::MutaliskParseVCFObj)

df.ref.mut.sigs
A data.frame of reference mutational signatures

target.mut.sigs
A character vector of target mutational signatures names
```

#### Value

A list with the following elements

- num.point.mutationsAn integer value count of total point mutations
- sub.typesA character vector of length 96
- sub.types.spectrumA numeric vector of length 96
- num.mut.sigsAn integer value (count of unique mutational signatures identified)
- · identified.mut.sigsA character vector where each element is a mutational signature identified
- identified.mut.sigs.probsA numeric vector where each element is the weight of mutational signature identified. The ordering follows identified.mut.sigs
- identified.mut.sigs.spectrumA numeric vector of length 96
- residuals A numeric vector of length 96
- rssA numeric value (residual sum of squares)
- cos.sim.scoreA numeric value (cosine similarity score between observed mutational spectrum and reconstructed mutational signatures)
- all.models.sigsA list where each element is a model; a model is a list of signatures identified)
- all.models.sigs.probsA list where each element is a model; a model is a list of contribution probabilities
- all.models.cos.sim.scoresA list where each element is a model; a model is a list of cosine similarity socres

RunMutPat

RunMutPat

## Description

Identifies mutational signatures using Mutational Patterns

#### Usage

```
RunMutPat(mut.pat.input, df.mut.pat.ref.sigs, target.mut.sigs,
  verbose = TRUE)
```

Sigmoid.Obj.Fn 53

#### **Arguments**

#### Value

A list with the following elements

- tumor.mutation.types.spectrumA numeric vector of length 96 'observed' spectrum
- identified.mutation.types.spectrumA numeric vector of length 96 'identified' spectrum
- residuals A numeric vector of length 96 residuals
- mutation.typesA character vector of length 96
- identified.mut.sigsA character vector where each element is a mutational signature identified
- identified.mut.sigs.contribution.weightsA numeric vector where each element is the weight of mutational signature identified. The ordering follows identified.mut.sigs
- cosine.similarity.scoreA numeric value

#### **Examples**

```
## Not run:
vcf.obj <- ReadVCF(vcf.file = "../data/sample/HNT-082-BT.final.call.vcf", genome = "hg19")
df.ref.mut.sigs <- GetPCAWGMutSigs()
target.mut.sigs <- GetPCAWGMutSigsNames()
RunMutPat(vcf.obj = vcf.obj,
df.ref.mut.sigs = df.ref.mut.sigs,
target.mut.sigs = target.mut.sigs)
## End(Not run)</pre>
```

Sigmoid.Obj.Fn

Sigmoid.Obj.Fn

#### **Description**

Sigmoid objective function

#### Usage

```
Sigmoid.Obj.Fn(C.refined, A.refined, C.artifactual, A.artifactual)
```

54 Test.Obj.Fn.1

## Arguments

C.refined A numeric value between 0 and 1.

A.refined A numeric value between 0 and 1.

C.artifactual A numeric value between 0 and 1.

A.artifactual A numeric value between 0 and 1.

#### Value

A numeric value between 0 and 1.

Test.Obj.Fn.1 Test.Obj.Fn.1

## Description

Test objective function 1

## Usage

```
Test.Obj.Fn.1(C.refined, A.refined, C.artifactual, A.artifactual)
```

## Arguments

C.refined A numeric value between 0 and 1.

A.refined A numeric value between 0 and 1.

C.artifactual A numeric value between 0 and 1.

A.artifactual A numeric value between 0 and 1.

#### Value

A numeric value between 0 and 1.

Test.Obj.Fn.2 55

Test.Obj.Fn.2

Test.Obj.Fn.2

## Description

Test objective function 2

## Usage

```
Test.Obj.Fn.2(C.refined, A.refined, C.artifactual, A.artifactual)
```

## Arguments

C.refined A numeric value between 0 and 1.

A.refined A numeric value between 0 and 1.

C.artifactual A numeric value between 0 and 1.

A.artifactual A numeric value between 0 and 1.

#### Value

A numeric value between 0 and 1.

```
TriNuc.Mutation.Type.Hex.Colors

**Constant**
```

## Description

Hex codes for the mutation types (for plotting purposes)

#### Usage

```
TriNuc.Mutation.Type.Hex.Colors
```

#### **Format**

An object of class character of length 6.

UpdateFilter

UpdateFilter

## Description

Update filter based on optim parameter values

## Usage

```
UpdateFilter(vcf.filter, param.values)
```

## Arguments

vcf.filter

A list from MakeFilterFromConfig

param.values

A numeric vector contains filtering value (same length with length(vcf.config.filter))

#### Value

Updated vcf.filter (list)

WriteFIREVATResultsToTSV

WriteFIREVATResultsToTSV

## Description

Writes FIREVAT results to a csv file

#### Usage

WriteFIREVATResultsToTSV(firevat.results)

## Arguments

firevat.results

List returned from RunFIREVAT

Write VCF 57

WriteVCF WriteVCF

## Description

Writes a vcf.obj to a .vcf file

## Usage

```
WriteVCF(vcf.obj, save.file)
```

## Arguments

```
vcf.obj (from the function ReadVCF) save.file (full path including filename)
```

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