

# Package ‘FIREVAT’

October 9, 2019

**Type** Package

**Title** FIREVAT, FInding REliable Variants without ArTifacts

**Description** FIREVAT is a variant filtering tool for cancer sequencing data, which uses mutational signatures to identify sequencing artifacts and low-quality variants.

**Version** 0.5.4

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**Depends** R (>= 3.5.0)

**Imports** data.table,  
 rngtools,  
 doRNG,  
 stringi,  
 bedr,  
 GA,  
 jsonlite,  
 yaml,  
 lsa,  
 ggpubr,  
 caTools,  
 ggrepel,  
 gridExtra,  
 ggplot2,  
 rmarkdown,  
 gtable,  
 dplyr,  
 extrafont,  
 IRanges,  
 BSgenome.Hsapiens.UCSC.hg19,  
 BSgenome.Hsapiens.UCSC.hg38,  
 MutationalPatterns,  
 deconstructSigs

**biocViews** CancerGenomics,  
 VariantRefinement,

VariantFiltering,  
 MutationalSignatures,  
 SomaticMutation,

**URL** <https://github.com/cgab-ncc/FIREVAT>

**Encoding** UTF-8

**LazyData** true

**RoxygenNote** 6.1.1

**Suggests** knitr

**VignetteBuilder** knitr

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AnnotateVCFObj	<i>AnnotateVCFObj</i>
----------------	-----------------------

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

<code>vcf.obj</code>	A list from <code>ReadVCF</code>
<code>bsg</code>	<code>BSgenome.Hsapiens.UCSC</code> object
<code>df.mut.pat.ref.sigs</code>	A data.frame from <code>MutPatParseRefMutSigs</code>
<code>target.mut.sigs</code>	A character vector of target mutational signatures from reference mutational signatures.
<code>sequencing.artifact.mut.sigs</code>	A character vector of sequencing artifact mutational signatures from reference mutational signatures.
<code>init.artifact.stop</code>	Numeric value less than 1. If the sum of sequencing artifact weights in <code>vcf.obj</code> is less than or equal to this value then this function returns <code>judgment = FALSE</code> , otherwise returns <code>judgment = TRUE</code> .
<code>verbose</code>	If <code>TRUE</code> , provides process detail. Default value is <code>TRUE</code> .

## Value

A list with the following elements

- `judgment`A boolean value
- `seq.art.sigs.weights.sum`A numeric value. Sum of sequencing artifact weights.

---

Chromosome.Names	<i>Constant</i>
------------------	-----------------

---

**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	<i>ComputeZScore</i>
---------------	----------------------

---

**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

---

ComputeZScoreEquiValue  
*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      *DecimalCeiling*

---

**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

---

Default.Obj.Fn	<i>Default.Obj.Fn</i>
----------------	-----------------------

---

### 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	<i>Transform default filtering parameters to a binary vector</i>
-----------------------	--

---

### 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 <a href="#">MakeFilter</a>
params.bit.len	A list with bit lengths of filtering parameters which is generated from <a href="#">ParameterToBits</a>

### Value

A binary vector



---

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 → C>A; 17:32 → C>G; 33:48 → C>T; 49:64 → T>A; 65:80 → T>C; 81:96 → 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)

### 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.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

<code>C.refined</code>	A numeric value between 0 and 1.
<code>A.refined</code>	A numeric value between 0 and 1.
<code>C.artifactual</code>	A numeric value between 0 and 1.
<code>A.artifactual</code>	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)`

### Arguments

<code>C.refined</code>	A numeric value between 0 and 1.
<code>A.refined</code>	A numeric value between 0 and 1.
<code>C.artifactual</code>	A numeric value between 0 and 1.
<code>A.artifactual</code>	A numeric value between 0 and 1.

**Value**

A numeric value between 0 and 1.

---

Exp.Weighted.A.Ref.Obj.Fn

*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.

---

Exp.Weighted.Obj.Fn.1    *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)

**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.Obj.Fn.2	<i>Exp.Weighted.Obj.Fn.2</i>
-----------------------	------------------------------

---

**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	<i>Exp.Weighted.Refined.Seq.Art.Only.Obj.Fn</i>
--	---

---

**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)

**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.

---

FilterByStrandBiasAnalysis
<i>FilterByStrandBiasAnalysis</i>

---

**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	<i>FilterVCF</i>
-----------	------------------

---

**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)
```

**Arguments**

<code>vcf.obj</code>	A list from <code>ReadVCF</code>
<code>vcf.filter</code>	A list from <code>MakeMuTect2Filter</code>
<code>config.obj</code>	A list from <code>ParseConfigFile</code>
<code>include.array</code>	A boolean vector
<code>force.include</code>	A boolean value. If TRUE, then uses 'include.array'
<code>verbose</code>	If true, provides process detail

**Value**

A list with the following elements

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

---

<code>GenerateConfigObj</code>	<i>Generate config.obj by checking vcf header</i>
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---

**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**

<code>vcf.obj</code>	A list from <a href="#">ReadVCF</a>
<code>save.config</code>	If true, save config.obj to config.path
<code>config.path</code>	File path to write config.obj (json or yaml)

**Value**

`config.obj`

---

GetCOSMICMutSigs	<i>GetCOSMICMutSigs</i>
------------------	-------------------------

---

**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

---

GetCOSMICMutSigsEtiologiesColors	<i>GetCOSMICMutSigsNames</i>
----------------------------------	------------------------------

---

**Description**  
Returns all COSMIC mutational signature etiologies and colors

**Usage**  
GetCOSMICMutSigsEtiologiesColors()

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

---

GetCOSMICMutSigsNames	<i>GetCOSMICMutSigsNames</i>
-----------------------	------------------------------

---

**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

<code>vcf.obj</code>	A list from ReadVCF
<code>bsg</code>	BSgenome.Hsapiens.UCSC object
<code>config.obj</code>	A list from ParseConfigFile
<code>lower.upper.list</code>	A list from GetParameterLowerUpperVector
<code>df.mut.pat.ref.sigs</code>	A data.frame from MutPatParseRefMutSigs
<code>target.mut.sigs</code>	A character vector of the target mutational signatures from reference mutational signatures.
<code>sequencing.artifact.mut.sigs</code>	A character vector of the sequencing artifact mutational signatures from reference mutational signatures.
<code>objective.fn</code>	Objective value derivation function.
<code>original.muts.seq.art.weights.sum</code>	A numeric value. 'seq.art.sigs.weights.sum' from CheckIfVariantRefinementIsNecessary
<code>ga.preemptive.killing</code>	If TRUE, then preemptively kills populations that yield greater sequencing artifact weights sum compared to the original mutational signatures analysis
<code>verbose</code>	If TRUE, provides process detail. Default value is TRUE.

## Value

A list with the following elements

- `judgment`A boolean value
- `seq.art.sigs.weights`A numeric value. Sum of sequencing artifact weights.

---

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 <a href="#">RunFIREVAT</a>
mutations.type	A string for type of mutations ('refined' or 'artifactual')
signatures	A string ('all', 'target', 'artifactual')

### Value

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

---

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)
```

### 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  $\max(\text{vcf.obj\$data}[[\text{param}]]) < 1$ , 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	<i>GetPCAWGMutSigs</i>
-----------------	------------------------

---

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

---

GetPCAWGMutSigsEtiologiesColors	<i>GetPCAWGMutSigsEtiologiesColors</i>
---------------------------------	--

---

Description

Returns the PCAWG mutational signatures etiologies and colors

Usage

```
GetPCAWGMutSigsEtiologiesColors()
```

Value

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

---

GetPCAWGMutSigsNames	<i>GetPCAWGMutSigsNames</i>
----------------------	-----------------------------

---

**Description**

Returns the PCAWG mutational signatures names

**Usage**

```
GetPCAWGMutSigsNames()
```

**Value**

a character vector of the PCAWG mutational signatures names

---

GetPCAWGPlatinumMutSigs	<i>GetPCAWGPlatinumMutSigs</i>
-------------------------	--------------------------------

---

**Description**

Returns the PCAWG platinum mutational signatures data

**Usage**

```
GetPCAWGPlatinumMutSigs()
```

**Value**

a data.frame of the PCAWG platinum mutatioanl signatures

---

GetPCAWGPlatinumMutSigsEtiologiesColors	<i>GetPCAWGPlatinumMutSigsEtiologiesColors</i>
---	--

---

**Description**

Returns the PCAWG platinum mutational signatures etiologies and colors

**Usage**

```
GetPCAWGPlatinumMutSigsEtiologiesColors()
```

**Value**

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

---

GetPCAWGPlatinumMutSigsNames
<i>GetPCAWGPlatinumMutSigsNames</i>

---

**Description**

Returns the PCAWG platinum mutational signatures names

**Usage**

GetPCAWGPlatinumMutSigsNames()

**Value**

a character vector of the PCAWG platinum mutational signatures names

---

GetSeedForVCF	<i>GetSeedForVCF</i>
---------------	----------------------

---

**Description**

Returns a seed integer based on VCF file size

**Usage**

GetSeedForVCF(vcf.file)

**Arguments**

vcf.file            (full path of a .vcf file)

**Details**

Returns the same seed integer for the same VCF file (based on file size)

**Value**

an integer value

---

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	<i>Leaky.ReLU.A.Ref.Obj.Fn</i>
-------------------------	--------------------------------

---

**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.

---

Leaky.ReLU.Obj.Fn	<i>Leaky.ReLU.Obj.Fn</i>
-------------------	--------------------------

---

**Description**

Lkey 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	<i>MakeFilter</i>
------------	-------------------

---

**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	<i>MutaliskParseVCFObj</i>
---------------------	----------------------------

---

**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	<i>MutPatParseRefMutSigs</i>
-----------------------	------------------------------

---

**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

---

MutPatParseVCFObj	<i>MutPatParseVCFObj</i>
-------------------	--------------------------

---

**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

---

ParameterToBits	<i>ParameterToBits</i>
-----------------	------------------------

---

**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  $\max(\text{vcf.obj\$data}[[\text{param}]]) < 1$ , then multiply multiplier to the vector

**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

---

ParseConfigFile	<i>ParseConfigFile</i>
-----------------	------------------------

---

**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.

---

PCAWG.Platinum.All.Technology.Related.Artifact.Signatures  
*Constant*

---

**Description**

PCAWG mutational signatures reported to be associated with sequencing artifacts

**Usage**

PCAWG.Platinum.All.Technology.Related.Artifact.Signatures

**Format**

An object of class character of length 9.

---

PCAWG.Possible.Sequencing.Artifact.Signatures  
*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.

---

PCAWG.Target.Mutational.Signatures  
*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")
```

**Arguments**

vcf.obj            [ReadVCF](#)  
ref.forward.strand.var  
                  A string value.  
ref.reverse.strand.var  
                  A string value.  
alt.forward.strand.var  
                  A string value.  
alt.reverse.strand.var  
                  A string value.  
perform.fdr.correction  
                  If TRUE, then performs false discovery rate correction  
fdr.correction.method  
                  A string value. FDR correction method (Refer to p.adjust() function)

**Value**

An updated vcf.obj

---

PlotMutaliskResults    *PlotMutaliskResults*

---

**Description**

Plots Mutalisk results

**Usage**

```
PlotMutaliskResults(mutalisk.results, signatures,  
  df.ref.sigs.groups.colors, 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  
`df.ref.sigs.groups.colors` A data.frame with signature groups and colors  
`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**

```
## Not run:
df.ref.mut.sigs <- GetPCAWGMutSigs()
target.mut.sigs <- GetPCAWGMutSigsNames()
vcf.obj <- ReadVCF(vcf.file = "../data/sample/P-233-CT.final.vcf")
mutalisk.results <- RunMutalisk(vcf.obj = vcf.obj,
                               df.ref.mut.sigs = df.ref.mut.sigs,
                               target.mut.sigs = target.mut.sigs)
p <- PlotMutaliskResults(mutalisk.results = mutalisk.results)
print(p)

## End(Not run)
```

---

PlotMutationTypes	<i>PlotMutationTypes</i>
-------------------	--------------------------

---

**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  
`title` Plot title  
`convert.to.percentage` if True convert y values to percentage (x 100); Default = T  
`show.legend` If True, show legend; 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:
p <- PlotMutationTypes(mutation.types = c("C>A", "C>G", "C>T", "T>A", "T>C", "T>G"),
  mutation.types.values = c(0.3, 0.3, 0.1, 0.1, 0.1, 0.1),
  mutation.types.colors = TriNuc.Mutation.Type.Hex.Colors,
  max.y.val = 0.5,
  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"))

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**

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

**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)
```

---

PlotTable	<i>PlotTable</i>
-----------	------------------

---

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	Padding size; Default = 20
font.size	Font size; Default = 14

Value

A plot

---

PlotTriNucSpectrum	<i>PlotTriNucSpectrum</i>
--------------------	---------------------------

---

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)
```

Arguments

sub.types	A character vector (types of 96 trinucleotide substitutions)
spectrum	A numeric vector (96 elements)
max.y.val	y axis maximum value
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	<i>PlotVCFStatsBoxPlots</i>
----------------------	-----------------------------

---

**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)
label.font.size	An integer value (label font size)
title.font.size	An integer value (title font size)

**Value**

A ggboxplot

---

PlotVCFStatsHistograms

*PlotVCFStatsHistograms*


---

## 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
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), "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	<i>PrepareAnnotationDB</i>
---------------------	----------------------------

---

**Description**

Prepares df.genes.of.interest from a vcf.obj ([ReadVCF](#)) of COSMIC or ClinVar vcf for [AnnotateVCFObj](#)

**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	<i>PrepareArtifactAnnotationTable</i>
--------------------------------	---------------------------------------

---

**Description**

Prepares artifactual mutations annotation (filtered, queried) table

**Usage**

PrepareArtifactAnnotationTable(data)

**Arguments**

data                      A list of elements returned from [RunFIREVAT](#)

**Value**

A data.frame

---

PrepareArtifactStrandBiasTable
<i>PrepareArtifactStrandBiasTable</i>

---

**Description**

Prepares artifactual mutations strand biased variants table

**Usage**

PrepareArtifactStrandBiasTable(data)

**Arguments**

data                    A list of elements returned from [RunFIREVAT](#)

**Value**

A data.frame

---

PrepareArtifactualMutsOptimizationIterationsPlot
<i>PrepareArtifactualMutsOptimizationIterationsPlot</i>

---

**Description**

Prepares artifactual mutations optimization iterations plot

**Usage**

PrepareArtifactualMutsOptimizationIterationsPlot(data)

**Arguments**

data                    A list of elements returned from [RunFIREVAT](#)

**Value**

A ggplot object

---

PrepareFilterCutoffsTable
<i>PrepareFilterCutoffsTable</i>

---

**Description**

Prepares filter cutoffs table for reporting

**Usage**

PrepareFilterCutoffsTable(data)

**Arguments**

data                    A list of elements returned from [RunFIREVAT](#)

**Value**

A data.frame

---

PrepareGeneticAlgorithmParametersTable
<i>PrepareGeneticAlgorithmParametersTable</i>

---

**Description**

Prepares Genetic Algorithm parameters table

**Usage**

PrepareGeneticAlgorithmParametersTable(data)

**Arguments**

data                    A list of elements returned from [RunFIREVAT](#)

**Value**

A data.frame

---

`PrepareIdentifiedSignaturesPlot`*PrepareIdentifiedSignaturesPlot*

---

**Description**

Prepares identified signatures plot for reporting

**Usage**

```
PrepareIdentifiedSignaturesPlot(data)
```

**Arguments**

`data`                      A list of elements returned from [RunFIREVAT](#)

**Value**

A ggarrange object

---

`PrepareMLEReconstructedSpectrumsPlot`*PrepareMLEReconstructedSpectrumsPlot*

---

**Description**

Prepares MLE reconstructed spectrums plot

**Usage**

```
PrepareMLEReconstructedSpectrumsPlot(data)
```

**Arguments**

`data`                      A list of elements returned from [RunFIREVAT](#)

**Value**

A ggarrange object



---

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  
*PrepareObservedSpectrumsPlot*

---

**Description**

Prepares observed spectrums plot

**Usage**

PrepareObservedSpectrumsPlot(data)

**Arguments**

data                    A list of elements returned from [RunFIREVAT](#)

**Value**

A ggarrange object

---

`PrepareOptimizationResultsTable`*PrepareOptimizationResultsTable*

---

**Description**

Prepares optimization results table

**Usage**

```
PrepareOptimizationResultsTable(data)
```

**Arguments**

`data`                      A list of elements returned from [RunFIREVAT](#)

**Value**

A data.frame

---

`PrepareOptimizedVCFStatisticsPlot`*PrepareOptimizedVCFStatisticsPlot*

---

**Description**

Prepares optimized VCF statistics plot

**Usage**

```
PrepareOptimizedVCFStatisticsPlot(data)
```

**Arguments**

`data`                      A list of elements returned from [RunFIREVAT](#)

**Value**

A ggarrange object

---

PrepareRefinedAnnotationTable
<i>PrepareRefinedAnnotationTable</i>

---

**Description**

Prepares refined mutations annotation (filtered, queried) table

**Usage**

PrepareRefinedAnnotationTable(data)

**Arguments**

data                    A list of elements returned from [RunFIREVAT](#)

**Value**

A data.frame

---

PrepareRefinedMutsOptimizationIterationsPlot
<i>PrepareRefinedMutsOptimizationIterationsPlot</i>

---

**Description**

Prepares refined mutations optimization iterations plot

**Usage**

PrepareRefinedMutsOptimizationIterationsPlot(data)

**Arguments**

data                    A list of elements returned from [RunFIREVAT](#)

**Value**

A ggplot object

---

`PrepareRefinedStrandBiasTable`*PrepareRefinedStrandBiasTable*

---

**Description**

Prepares refined mutations strand biased variants table

**Usage**

```
PrepareRefinedStrandBiasTable(data)
```

**Arguments**

`data`                      A list of elements returned from [RunFIREVAT](#)

**Value**

A data.frame

---

`PrepareResidualSpectrumsPlot`*PrepareResidualSpectrumsPlot*

---

**Description**

Prepares residual spectrums plot

**Usage**

```
PrepareResidualSpectrumsPlot(data)
```

**Arguments**

`data`                      A list of elements returned from [RunFIREVAT](#)

**Value**

A ggarrange object

---

PrepareTrinucleotideSpectrumsTable
<i>PrepareTrinucleotideSpectrumsTable</i>

---

**Description**

Prepares trinucleotide spectrums table

**Usage**

PrepareTrinucleotideSpectrumsTable(data)

**Arguments**

data                   A list of elements returned from [RunFIREVAT](#)

**Value**

A data.frame

---

PrintLog	<i>PrintLog</i>
----------	-----------------

---

**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.

---

QueryAnnotatedVCF	<i>FilterAnnotatedVCF</i>
-------------------	---------------------------

---

**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`  
[AnnotateVCFObj](#)
- `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
<i>ReadOptimizationIterationReport</i>

---

**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	<i>ReadVCF</i>
---------	----------------

---

**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.info 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'

---

ReportFIREVATResults	<i>ReportFIREVATResults</i>
----------------------	-----------------------------

---

**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 meta-data related to the refinement process.

**Usage**

```
RunFIREVAT(vcf.file, vcf.file.genome = "hg19", config.file,
  df.ref.mut.sigs = GetPCAWGMutSigs(),
  target.mut.sigs = GetPCAWGMutSigsNames(),
  df.ref.mut.sigs.groups.colors = GetPCAWGMutSigsEtiologiesColors(),
  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,
  ga.seed = NULL, 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,
  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 either 'hg19' or 'hg38'.
config.file	String value corresponding to input configuration file. For more details please refer to ...



<code>df.ref.mut.sigs</code>	A data.frame of the reference mutational signatures
<code>target.mut.sigs</code>	A character vector of the target mutational signatures from reference mutational signatures.
<code>df.ref.mut.sigs.groups.colors</code>	A data.frame of the reference mutational signatures groups and colors
<code>sequencing.artifact.mut.sigs</code>	A character vector of the sequencing artifact mutational signatures from reference mutational signatures.
<code>num.cores</code>	Number of cores to allocate
<code>output.dir</code>	String value of the desired output directory
<code>mode</code>	String value. The value should be either 'ga' or 'manual'.
<code>init.artifact.stop</code>	Numeric value less than 1. If the sum of sequencing artifact weights in the user-specified original VCF file (i.e. <code>vcf.file</code> ) 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'.
<code>objective.fn</code>	Objective value derivation function. Default: <code>Default.Obj.Fn</code> .
<code>use.suggested.soln</code>	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.
<code>ga.type</code>	String value. The value should be either 'binray' or 'real-valued'.
<code>ga.pop.size</code>	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.
<code>ga.max.iter</code>	Integer value of the Genetic Algorithm 'maximum iterations' parameter. Default: 100. This value should be set based on the number of filter parameters. Recommendation: same as 'ga.pop.size'.
<code>ga.run</code>	Integer value of the Genetic Algorithm 'run' parameter. Default: 50. This value should be set based on the 'ga.max.iter' parameter. Recommendation: 25 percent of 'ga.max.iter'.
<code>ga.pmutation</code>	Float value of the Genetic Algorithm 'mutation probability' parameter. Default: 0.1.
<code>ga.preemptive.killing</code>	If TRUE, then preemptively kills populations that yield greater sequencing artifact weights sum compared to the original mutational signatures analysis
<code>ga.seed</code>	Integer value of the Genetic Algorithm 'seed' parameter. Default: NULL.
<code>mutalisk</code>	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,

`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	<i>RunGAMode</i>
-----------	------------------

---

**Description**

Runs FIREVAT ga mode

**Usage**

RunGAMode(data)

**Arguments**

data                    A list from RunFIREVAT

**Value**

A list

---

RunManualMode	<i>RunManualMode</i>
---------------	----------------------

---

**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**

<code>vcf.obj</code>	A list (from <code>firevat_vcf::ReadVCF</code> )
<code>df.ref.mut.sigs</code>	A data.frame of reference mutational signatures
<code>target.mut.sigs</code>	A character vector of target mutational signatures names to identify from
<code>random.sampling.candidate.mut.sigs</code>	A character vector of mutational signatures names that gets appended to the list of candidate mutational signatures so that these are always considered.
<code>method</code>	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
<code>n.sample</code>	An integer value ('random.sampling' method parameter) Number of signatures to choose for each iteration of random sampling).
<code>n.iter</code>	An integer value ('random.sampling' method parameter). Number of iterations to perform random sampling.
<code>verbose</code>	If true, provides process details

**Value**

A list with the following elements

- `num.point.mutations` An integer value - count of total point mutations
- `sub.types` A character vector of length 96
- `sub.types.spectrum` A numeric vector of length 96
- `num.mut.sigs` An integer value (count of unique mutational signatures identified)
- `identified.mut.sigs` A character vector where each element is a mutational signature identified
- `identified.mut.sigs.probs` A 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 scores

---

RunMutaliskHelper

*RunMutaliskHelper*


---

## Description

Helper function for RunMutalisk

## Usage

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

## 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

- 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 scores

RunMutPat

*RunMutPat*

## Description

Identifies mutational signatures using Mutational Patterns

## Usage

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

## Arguments

mut.pat.input    A list from [MutPatParseVCFObj](#)  
df.mut.pat.ref.sigs    A data.frame returned by [MutPatParseRefMutSigs](#)  
target.mut.sigs    A character vector of target mutational signatures names  
verbose    If true, provides process details

## 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
- residualsA 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)
```

---

Sigmoid.Obj.Fn	<i>Sigmoid.Obj.Fn</i>
----------------	-----------------------

---

Description

Sigmoid objective function

Usage

```
Sigmoid.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.

---

Test.Obj.Fn.1	<i>Test.Obj.Fn.1</i>
---------------	----------------------

---

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	<i>Test.Obj.Fn.2</i>
---------------	----------------------

---

**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
<i>Constant</i>

---

**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	<i>UpdateFilter</i>
--------------	---------------------

---

**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
<i>WriteFIREVATResultsToTSV</i>

---

**Description**

Writes FIREVAT results to a csv file

**Usage**

WriteFIREVATResultsToTSV(firevat.results)

**Arguments**

- firevat.results      List returned from RunFIREVAT

---

`WriteVCF`*WriteVCF*

---

**Description**

Writes a vcf.obj to a .vcf file

**Usage**

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

**Arguments**

<code>vcf.obj</code>	(from the function <code>ReadVCF</code> )
<code>save.file</code>	(full path including filename)

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