# Package 'ICAMS'

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Type Package

```
Title In-depth Characterization and Analysis of Mutational Signatures ('ICAMS')
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Description Analysis and visualization of experimentally elucidated mutational
      signatures -- the kind of analysis and visualization in Boot et al.,
      "In-depth characterization of the cisplatin mutational signature in
      human cell lines and in esophageal and liver tumors", Genome Research 2018,
      <doi:10.1101/gr.230219.117>. 'ICAMS' stands for In-depth Characterization
      and Analysis of Mutational Signatures. 'ICAMS' has functions to read in
      variant call files (VCFs) and to collate the corresponding catalogs of
      mutational spectra and to analyze and plot catalogs of mutational spectra
      and signatures. Handles both ``counts-based" and ``density-based" catalogs
      of mutational spectra or signatures.
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```

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stringi,
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	abundance <i>K-mer abundances</i> .	

# **Description**

An R list with one element each for BSgenome.Hsapiens.1000genomes.hs37d5, BSgenome.Hsapiens.UCSC.hg38 and BSgenome.Mmusculus.UCSC.mm10. Each element is in turn a sub-list keyed by exome, transcript, and genome. Each element of the sub list is keyed by the number of rows in the catalog class (as a string, e.g. "78", not 78). The keys are: 78 (DBS78Catalog), 96 (SBS96Catalog), 136 (DBS136Catalog), 144 (DBS144Catalog), 192 (SBS192Catalog), and 1536 (SBS1536Catalog). So, for example to get the exome abundances for SBS96 catalogs for BSgenome.Hsapiens.UCSC.hg38 exomes one would reference

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```
all.abundance[["BSgenome.Hsapiens.UCSC.hg38"]][["exome"]]["96"] or all.abundance$BSgenome.Hsapiens.UCSC.hg38$exome$"96". The value of the abundance is an integer vector with the K-mers as names and each value being the count of that K-mer.
```

### Usage

all.abundance

#### **Format**

See Description.

# **Examples**

```
all.abundance$BSgenome.Hsapiens.UCSC.hg38$transcript$`144`

# AA AC AG AT CA CC ...

# 90769160 57156295 85738416 87552737 83479655 63267896 ...

# There are 90769160 AAs on the sense strands of transcripts in # this genome.
```

as.catalog

Create a catalog from a numeric matrix or numeric data. frame.

# **Description**

Create a catalog from a numeric matrix or numeric data. frame.

# Usage

```
as.catalog(object, ref.genome = NULL, region = "unknown",
  catalog.type = "counts", abundance = NULL)
```

# **Arguments**

object A numeric matrix or numeric data. frame. This object must have rownames to

denote the mutation types. See CatalogRowOrder for more details.

ref.genome A ref.genome argument as described in ICAMS.

region A character string designating a region, one of genome, transcript, exome,

unknown; see ICAMS.

catalog.type One of "counts", "density", "counts.signature", "density.signature".

abundance If NULL, then inferred if ref.genome is one of the reference genomes known

to ICAMS and region is not unknown. See ICAMS. The argument abundance should contain the counts of different source sequences for mutations in the

same format as the numeric vectors in all.abundance.

# Value

A catalog as described in ICAMS.

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### **Examples**

CatalogRowOrder

Standard order of row names in a catalog.

# **Description**

This data is designed for those who need to create their own catalogs from formats not supported by this package. The rownames denote the mutation types. For example, for SBS96 catalogs, the rowname AGAT represents a mutation from AGA > ATA.

# Usage

```
catalog.row.order
```

#### **Format**

A list of character vectors indicating the standard orders of row names in catalogs.

### Note

In ID (insertion and deletion) catalogs, deletion repeat sizes range from 0 to 5+, but for plotting and end-user documentation deletion repeat sizes range from 1 to 6+.

# **Examples**

```
catalog.row.order$SBS96
# "ACAA" "ACCA" "ACCA" "CCCA" "CCCA" "CCCA" "CCTA" ...
# There are altogether 96 row names to denote the mutation types
# in SBS96 catalog.
```

CollapseCatalog

"Collapse" a catalog.

# **Description**

- 1. Take a mutational spectrum or signature catalog that is based on a fined-grained set of features (for example, single-nucleotide substitutions in the context of the preceding and following 2 bases).
- 2. Collapse it to a catalog based on a coarser-grained set of features (for example, single-nucleotide substitutions in the context of the immediately preceding and following bases).

Collapse192CatalogTo96 Collapse an SBS 192 catalog to an SBS 96 catalog.

Collapse1536CatalogTo96 Collapse an SBS 1536 catalog to an SBS 96 catalog.

Collapse144CatalogTo78 Collapse a DBS 144 catalog to a DBS 78 catalog.

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### Usage

```
Collapse192CatalogTo96(catalog)
Collapse1536CatalogTo96(catalog)
Collapse144CatalogTo78(catalog)
```

# **Arguments**

catalog A catalog as defined in ICAMS.

#### Value

A catalog as defined in ICAMS.

# **Examples**

FindDelMH

Return the length of microhomology at a deletion.

# **Description**

Return the length of microhomology at a deletion.

### Usage

```
FindDelMH(context, deleted.seq, pos, trace = 0)
```

# **Arguments**

context The deleted sequence plus ample surrounding sequence on each side (at least as

long as del. sequence).

deleted.seq The deleted sequence in context.

pos The position of del. sequence in context.

trace If > 0, cat various messages.

# **Details**

This function is primarily for internal use, but we export it to document the underlying logic.

Example:

GGCTAGTT aligned to GGCTAGAACTAGTT with a deletion represented as:

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```
GGCTAGAACTAGTT
GG-----CTAGTT GGCTAGTT GG[CTAGAA]CTAGTT
---- ----
```

Presumed repair mechanism leading to this:

GGCTAGAACTAGTT

=>

GGCTAG TT

=>

GGCTAGTT CCGATCAA

Variant-caller software can represent the same deletion in several different, but completely equivalent, ways.

```
GGCTAGTT GGCTAGTT GGC[TAGAAC]TAGTT

* --- * ---

GGCT-----AGTT GGCTAGTT GGCT[AGAACT]AGTT

** -- ** --

GGCTA-----GTT GGCTAGTT GGCTA[GAACTA]GTT

*** - *** -

GGCTAG-----TT GGCTAGTT GGCTAG[AACTAG]TT
```

A deletion in a *repeat* can also be represented in several different ways. A deletion in a repeat is abstractly equivalent to microhomology that spans the entire deleted sequence. For example;

```
GACTAGCTAGTT
GACTAGTT GACTA[GCTA]GTT

*** -*** -

is really a repeat

GACTAG----TT GACTAGTT GACTAG[CTAG]TT

**** ----

GACT----AGTT GACTAGTT GACT[AGCT]AGTT

** --** --
```

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This function only flags this case with a -1 return; it does not figure out the repeat extent.

This function finds:

1. The maximum match of undeleted sequence to the left of the deletion that is identical to the right end of the deleted sequence, and

2. The maximum match of undeleted sequence to the right of the deletion that is identical to the left end of the deleted sequence.

The microhomology sequence is the concatenation of items (1) and (2).

# Value

The length of the maximum microhomology of del. sequence in context.

# **Examples**

```
# GAGAGG[CTAGAA]CTAGTT
# ---- FindDelMH("GGAGAGGCTAGAACTAGTTAAAAA", "CTAGAA", 8, trace = 0) # 4
```

GetVAF

Extract the VAFs (variant allele frequencies) from a VCF file.

### **Description**

Extract the VAFs (variant allele frequencies) from a VCF file.

# Usage

```
GetStrelkaVAF(vcf)
GetMutectVAF(vcf)
```

### **Arguments**

vcf

Said VCF as a data.frame.

### Value

A vector of VAFs, one for each row of vcf.

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ICAMS: In-depth Characterization and Analysis of Mutational Signatures

### **Description**

Analysis and visualization of experimentally elucidated mutational signatures – the kind of analysis and visualization in Boot et al., "In-depth characterization of the cisplatin mutational signature in human cell lines and in esophageal and liver tumors",

Genome Research 2018, https://doi.org/10.1101/gr.230219.117. "ICAMS" stands for In-depth Characterization and Analysis of Mutational Signatures. ICAMS has functions to read in variant call files (VCFs) and to collate the corresponding catalogs of mutational spectra and to analyze and plot catalogs of mutational spectra and signatures. Handles both "counts-based" and "density-based" catalogs of mutational spectra or signatures.

#### **Details**

ICAMS can read in VCFs generated by Strelka or Mutect, and collate the mutations into "catalogs" of mutational spectra. ICAMS can create and plot catalogs of mutational spectra or signatures for single base substitutions (SBS), double base substitutions (DBS), and small insertions and deletions (ID). It can also read and write these catalogs.

### **Catalogs**

A key data type in ICAMS is a "catalog" of mutation counts, of mutation densities, or of mutational signatures.

Catalogs are R S3 objects of class matrix and one of several additional classes that specify the types of the mutations represented in the catalog (e.g. SBS96, ID, etc, ...). The possible additional classes are one of SBS96Catalog, SBS192Catalog, SBS1536Catalog, DBS78Catalog, DBS144Catalog, DBS136Catalog, IndelCatalog. as.catalog is the main constructor.

Conceptually, a catalog has one of the following types, which are indicated in the attribute catalog. type:

- 1. Matrix of mutation counts (one column per sample), representing (counts-based) mutational spectra (catalog.type = "counts").
- 2. Matrix of mutation densities, i.e. mutations per occurrences of source sequences (one column per sample), representing (density-based) mutational spectra (catalog.type = "density").
- 3. Matrix of mutational signatures, which are similar to spectra. However where spectra consist of counts or densities of mutations in each mutation class (e.g. ACA > AAA, ACA > AGA, ACA > ATA, ACC > AAC, ...), signatures consist of the proportions of mutations in each class (with all the proportions summing to 1). A mutational signature can be based on either:
  - mutation counts (a "counts-based mutational signature", catalog.type = "counts.signature"), or
  - $\bullet \ \ mutation \ densities \ (a \ "density-based \ mutational \ signature", \ catalog. \ type = "density.signature").$

Catalogs also have the attribute abundance, which contains the counts of different source sequences for mutations. For example, for SBSs in trinucleotide context, the abundances would be the counts of each trinucleotide in the human genome, exome, or in the transcribed region of the genome. See below under TransformCatalog for more information. Abundances logically depend on the species in question and on the part of the genome being analyzed.

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In ICAMS functions these can sometimes be inferred from catalog class attribute and function arguments region, ref.genome, and catalog.type. Otherwise they can be provided as an abundance argument. See all.abundance for examples.

Possible values for region are the strings genome, transcript, exome, and unknown; transcript includes entire transcribed regions, i.e. the introns as well as the exons.

If you need to create a catalog from a source other than this package (i.e. other than with ReadCatalog or StrelkaSBSVCFFilesToCatalog, MutectVCFFilesToCatalog, etc.), then use as.catalog.

### **Creating catalogs from variant call files (VCF files)**

- 1. StrelkaSBSVCFFilesToCatalog creates 3 SBS catalogs (96, 192, 1536) and 3 DBS catalogs (78, 136, 144) from the Strelka SBS VCFs.
- 2. StrelkaIDVCFFilesToCatalog creates ID (indel) catalog from the Strelka ID VCFs.
- 3. MutectVCFFilesToCatalog creates 3 SBS catalogs (96, 192, 1536), 3 DBS catalogs (78, 136, 144) and ID (indel) catalog from the Mutect VCFs.

### Plotting catalogs

The PlotCatalog functions plot mutational spectra for **one** sample or plot **one** mutational signature.

The PlotCatalogToPdf functions plot catalogs of mutational spectra or of mutational signatures to a PDF file.

### Wrapper functions to create catalogs from VCFs and plot the catalogs to PDF files

- 1. StrelkaSBSVCFFilesToCatalogAndPlotToPdf creates all type of SBS and DBS catalogs from Strelka SBS VCFs and plots the catalogs.
- 2. StrelkaIDVCFFilesToCatalogAndPlotToPdf creates an ID (indel) catalog from Strelka ID VCFs and plot it.
- 3. MutectVCFFilesToCatalogAndPlotToPdf creates all types of SBS, DBS, and ID catalogs from Mutect VCFs and plots the catalogs.

### The ref. genome (reference genome) argument

Many functions take the argument ref.genome.

In order to create a mutational spectrum catalog, ICAMS needs to know the sequence context of the mutations in the VCF file. For this, ICAMS needs the reference genome sequence that matches the VCF file. The ref.genome argument provides this.

ref.genome can be either

- 1. A variable from the Bioconductor BSgenome package that contains a particular reference genome, for example BSgenome. Hsapiens. 1000genomes. hs37d5.
- 2. The strings "hg38" or "GRCh38" are shorthand for BSgenome.Hsapiens.UCSC.hg38, the strings "hg19" or "GRCh37" are shorthand for BSgenome.Hsapiens.1000genomes.hs37d5, and the strings "mm10" or "GRCm38" are shorthand for BSgenome.Mmusculus.UCSC.mm10.

The Bioconductor BSgenome package, two human genomes BSgenome are "imported" by ICAMS and therefore should be installed when ICAMS is installed. These genomes are:

- BSgenome.Hsapiens.1000genomes.hs37d5
- BSgenome.Hsapiens.UCSC.hg38

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Any other needed reference genomes must be installed separately by the user. Use available.genomes() to get the list of available genomes. Further instructions are at

https://bioconductor.org/packages/release/bioc/html/BSgenome.html.

Use of ICAMS with other reference genomes is restricted to catalog. type of counts or counts. signature unless the user also creates the necessary abundance vectors. See all.abundance.

### Writing catalogs to files

The WriteCatalog functions write a catalog to a file.

### Reading catalogs

The ReadCatalog functions read a file that contains a catalog in standardized format.

### **Transforming catalogs**

The TransformCatalog function transforms catalogs of mutational spectra or signatures to account for differing abundances of the source sequence of the mutations in the genome.

For example, mutations from ACG are much rarer in the human genome than mutations from ACC simply because CG dinucleotides are rare in the genome. Consequently, there are two possible representations of mutational spectra or signatures. One representation is based on mutation counts as observed in a given genome or exome, and this approach is widely used, as, for example, at https://cancer.sanger.ac.uk/cosmic/signatures, which presents signatures based on observed mutation counts in the human genome. We call these "counts-based spectra" or "counts-based signatures".

Alternatively, mutational spectra or signatures can be represented as mutations per source sequence, for example the number of ACT > AGT mutations occurring at all ACT 3-mers in a genome. We call these "density-based spectra" or "density-based signatures".

This function can also transform spectra based on observed genome-wide counts to "density"-based catalogs. In density-based catalogs mutations are expressed as mutations per source sequences. For example, a density-based catalog represents the proportion of ACCs mutated to ATCs, the proportion of ACGs mutated to ATGs, etc. This is different from counts-based mutational spectra catalogs, which contain the number of ACC > ATC mutations, the number of ACG > ATG mutations, etc.

This function can also transform observed-count based spectra or signatures from genome to exome based counts, or between different species (since the abundances of source sequences vary between genome and exome and between species).

# Collapsing catalogs

The CollapseCatalog functions

- 1. Take a mutational spectrum or signature catalog that is based on a fined-grained set of features (for example, single-nucleotide substitutions in the context of the preceding and following 2 bases).
- 2. Collapse it to a catalog based on a coarser-grained set of features (for example, single-nucleotide substitutions in the context of the immediately preceding and following bases).

# Data

1. CatalogRowOrder Standard order of rownames in a catalog. The rownames encode the type of each mutation. The rownames denote the mutation types. For example, for SBS96 catalogs, the rowname AGAT represents a mutation from AGA > ATA.

2. TranscriptRanges Transcript ranges and strand information for a particular reference genome.

MutectVCFFilesToCatalog

Create SBS, DBS and Indel catalogs from Mutect VCF files

# **Description**

Create 3 SBS catalogs (96, 192, 1536), 3 DBS catalogs (78, 136, 144) and Indel catalog from the Mutect VCFs specified by files

### Usage

```
MutectVCFFilesToCatalog(files, ref.genome, trans.ranges = NULL,
  region = "unknown")
```

# **Arguments**

ref.genome

files Character vector of file paths to the Mutect VCF files. A ref. genome argument as described in ICAMS.

trans.ranges a data.table which contains transcript range and strand information. Please

refer to TranscriptRanges for more details.

A character string designating a genomic region; see as. catalog and ICAMS. region

# **Details**

This function calls VCFsToSBSCatalogs, VCFsToDBSCatalogs and VCFsToIDCatalogs

### Value

A list of 3 SBS catalogs (one each for 96, 192, and 1536), 3 DBS catalogs (one each for 78, 136, and 144) and ID catalog. If trans.ranges = NULL, SBS 192 and DBS 144 catalog will not be generated. Each catalog has attributes added. See as.catalog for more details.

# Note

SBS 192 and DBS 144 catalogs include only mutations in transcribed regions. In ID (insertion and deletion) catalogs, deletion repeat sizes range from 0 to 5+, but for plotting and end-user documentation deletion repeat sizes range from 1 to 6+.

```
file <- c(system.file("extdata",</pre>
                        "Mutect.GRCh37.vcf",
                       package = "ICAMS"))
catalogs <- MutectVCFFilesToCatalog(file, ref.genome = "hg19",</pre>
                                       trans.ranges = trans.ranges.GRCh37,
                                       region = "genome")
```

 ${\tt MutectVCFFilesToCatalogAndPlotToPdf}$ 

Create SBS, DBS and Indel catalogs from Mutect VCF files and plot them to PDF

## **Description**

Create 3 SBS catalogs (96, 192, 1536), 3 DBS catalogs (78, 136, 144) and Indel catalog from the Mutect VCFs specified by files and plot them to PDF

### Usage

```
MutectVCFFilesToCatalogAndPlotToPdf(files, ref.genome,
  trans.ranges = NULL, region = "unknown", output.file)
```

### **Arguments**

files Character vector of file paths to the Mutect VCF files. ref.genome A ref. genome argument as described in ICAMS. trans.ranges

a data.table which contains transcript range and strand information. Please

refer to TranscriptRanges for more details.

region A character string designating a genomic region; see as . catalog and ICAMS.

The name of the PDF file to be produced. output.file

### **Details**

This function calls MutectVCFFilesToCatalog and PlotCatalogToPdf

### Value

A list of 3 SBS catalogs (one each for 96, 192, and 1536), 3 DBS catalogs (one each for 78, 136, and 144), Indel catalog and their graphs plotted to PDF with specified file name. If trans.ranges = NULL, SBS 192 and DBS 144 catalog will not be generated and plotted. Each catalog has attributes added. See as.catalog for more details.

# Note

SBS 192 and DBS 144 catalogs include only mutations in transcribed regions. In ID (insertion and deletion) catalogs, deletion repeat sizes range from 0 to 5+, but for plotting and end-user documentation deletion repeat sizes range from 1 to 6+.

```
file <- c(system.file("extdata",</pre>
                       "Mutect.GRCh37.vcf",
                       package = "ICAMS"))
catalogs <-
  MutectVCFFilesToCatalogAndPlotToPdf(file, ref.genome = "hg19",
                                        trans.ranges = trans.ranges.GRCh37,
                                        region = "genome",
                                        output.file = file.path(tempdir(), "Mutect.pdf"))
```

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PlotCatalog	Plot one spectrum or signature.	

# **Description**

Plot the spectrum of **one** sample or plot **one** signature. The type of graph is based on one attribute("catalog.type") of the input catalog. You can first use TransformCatalog to get different types of catalog and then do the plotting.

# Usage

```
PlotCatalog(catalog, plot.SBS12 = NULL, cex = NULL, grid = NULL,
    upper = NULL, xlabels = NULL)
```

# **Arguments**

catalog	A catalog as defined in ICAMS with attributes added. See as.catalog for more details.
plot.SBS12	Only meaningful for class SBS192Catalog; if TRUE, generate an abbreviated plot of only SBS without context, i.e. C>A, C>G, C>T, T>A, T>C, T>G each on transcribed and untranscribed strands, rather than SBS in trinucleotide context, e.g. ACA > AAA, ACA > AGA,, TCT > TAT,
cex	A numerical value giving the amount by which mutation class labels, mutation counts(if it exists), y axis and its labels, x axis labels and its annotations(if it exists), sample name and legend(if it exists) should be magnified relative to the default. Only implemented for SBS96Catalog, SBS192Catalog and DBS144Catalog.
grid	A logical value indicating whether to draw grid lines. Only implemented for SBS96Catalog.
upper	A logical value indicating whether to draw horizontal lines and the names of major mutation class on top of graph. Only implemented for SBS96Catalog.
xlabels	A logical value indicating whether to draw x axis labels. Only implemented for SBS96Catalog.

## Value

```
invisible(TRUE)
```

# Note

The sizes of repeats involved in deletions range from 0 to 5+ in the mutational-spectra and signature catalog rownames, but for plotting and end-user documentation deletion repeat sizes range from 1 to 6+.

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```
catalog.type = "counts")
colnames(catSBS96) <- "sample"
PlotCatalog(catSBS96)</pre>
```

 ${\tt PlotCatalogToPdf}$ 

Plot catalog to a PDF file.

# Description

Plot catalog to a PDF file. The type of graph is based on one attribute("catalog.type") of the input catalog. You can first use TransformCatalog to get different types of catalog and then do the plotting.

# Usage

```
PlotCatalogToPdf(catalog, file, plot.SBS12 = NULL, cex = NULL,
   grid = NULL, upper = NULL, xlabels = NULL)
```

# **Arguments**

catalog	A catalog as defined in ICAMS with attributes added. See as.catalog for more details.
file	The name of the PDF file to be produced.
plot.SBS12	Only meaningful for class SBS192Catalog; if TRUE, generate an abbreviated plot of only SBS without context, i.e. C>A, C>G, C>T, T>A, T>C, T>G each on transcribed and untranscribed strands, rather than SBS in trinucleotide context, e.g. ACA > AAA, ACA > AGA,, TCT > TAT, There are 12 bars in the graph.
cex	A numerical value giving the amount by which mutation class labels, mutation counts(if it exists), y axis and its labels, x axis labels and its annotations(if it exists), sample name and legend(if it exists) should be magnified relative to the default. Only implemented for SBS96Catalog, SBS192Catalog and DBS144Catalog.
grid	A logical value indicating whether to draw grid lines. Only implemented for SBS96Catalog.
upper	A logical value indicating whether to draw horizontal lines and the names of major mutation class on top of graph. Only implemented for SBS96Catalog.
xlabels	A logical value indicating whether to draw x axis labels. Only implemented for SBS96Catalog.

### Value

```
invisible(TRUE)
```

# Note

The sizes of repeats involved in deletions range from 0 to 5+ in the mutational-spectra and signature catalog rownames, but for plotting and end-user documentation deletion repeat sizes range from 1 to 6+.

### **Examples**

ReadAndSplitMutectVCFs

Read and split Mutect VCF files.

# **Description**

Read and split Mutect VCF files.

### Usage

ReadAndSplitMutectVCFs(files)

### **Arguments**

files

Character vector of file paths to the Mutect VCF files.

# Value

A list with 3 in-memory VCFs and two left-over VCF-like data frames with rows that were not incorporated into the first 3 VCFs, as follows:

- 1. SBS VCF with only single base substitutions.
- 2. DBS VCF with only doublet base substitutions as called by Mutect.
- 3. ID VCF with only small insertions and deletions.
- 4. other.subs VCF like data.frame with rows for coordinate substitutions involving 3 or more nucleotides, e.g. ACT > TGA or AACT > GGTA.
- 5. multiple.alternative.alleles VCF like data.frame with rows for variants with multiple alternative alleles, for example ACT mutated to both AGT and ACT at the same position.

### See Also

MutectVCFFilesToCatalog

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```
Read And Split Strelka SBSVCFs \\
```

Read and split Strelka SBS VCF files.

# Description

Read and split Strelka SBS VCF files.

### Usage

```
ReadAndSplitStrelkaSBSVCFs(files)
```

# **Arguments**

files

Character vector of file paths to the Strelka SBS VCF files.

# Value

A list of 3 in-memory objects as follows:

- 1. SBS.vcfs List of data.frames of pure SBS mutations no DBS or 3+BS mutations.
- 2. DBS.vcfs List of data.frames of pure DBS mutations no SBS or 3+BS mutations.
- 3. ThreePlus List of data.tables with the key CHROM, LOW.POS, HIGH.POS. containing rows that that in the input that did not represent SBSs or DBSs.

### See Also

```
{\tt StrelkaSBSVCFFilesToCatalog}
```

# **Examples**

ReadCatalog

Read catalog.

# **Description**

Read a catalog in standardized format from path.

```
ReadCatalog(file, ref.genome, region, catalog.type, strict = TRUE)
```

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# **Arguments**

file Path to a catalog on disk in the standardized format.

ref.genome A ref.genome argument as described in ICAMS.

region region A character string designating a genomic region; see as.catalog and

ICAMS.

catalog.type One of "counts", "density", "counts.signature", "density.signature".

strict If TRUE, do additional checks on the input, and stop if the checks fail.

#### **Details**

See also WriteCatalog

#### Value

A catalog as an S3 object; see as.catalog.

#### Note

In ID (insertion and deletion) catalogs, deletion repeat sizes range from 0 to 5+, but for plotting and end-user documentation deletion repeat sizes range from 1 to 6+.

# **Examples**

ReadStrelkaIDVCFs

Read Strelka ID (insertion and deletion) VCF files.

# Description

Read Strelka ID (insertion and deletion) VCF files.

# Usage

```
ReadStrelkaIDVCFs(files)
```

# Arguments

files Character vector of file paths to the VCF files.

# Value

A list of vcfs from files.

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### Note

In ID (insertion and deletion) catalogs, deletion repeat sizes range from 0 to 5+, but for plotting and end-user documentation deletion repeat sizes range from 1 to 6+.

# See Also

```
StrelkaIDVCFFilesToCatalog
```

### **Examples**

revc

Reverse complement every string in string.vec.

# Description

```
Based on reverseComplement. Handles IUPAC ambiguity codes but not "u" (uracil). (see <a href="https://en.wikipedia.org/wiki/Nucleic_acid_notation">https://en.wikipedia.org/wiki/Nucleic_acid_notation</a>).
```

# Usage

```
revc(string.vec)
```

# **Arguments**

string.vec

A character vector.

# Value

A character vector with the reverse complement of every string in string.vec.

```
revc("aTgc") # GCAT

# A vector and strings with ambiguity codes
revc(c("ATGC", "aTGc", "wnTCb")) # GCAT GCAT VGANW

## Not run:
revc("ACGU") # An error
## End(Not run)
```

#### StrelkaIDVCFFilesToCatalog

Create ID (indel) catalog from Strelka ID VCF files

# **Description**

Create ID (indel) catalog from the Strelka ID VCFs specified by files

### Usage

```
StrelkaIDVCFFilesToCatalog(files, ref.genome, region = "unknown")
```

### **Arguments**

files Character vector of file paths to the Strelka ID VCF files.

ref.genome A ref.genome argument as described in ICAMS.

region A character string designating a genomic region; see as.catalog and ICAMS.

### **Details**

This function calls VCFsToIDCatalogs

### Value

An ID (indel) catalog with attributes added. See as.catalog for more details.

# Note

In ID (insertion and deletion) catalogs, deletion repeat sizes range from 0 to 5+, but for plotting and end-user documentation deletion repeat sizes range from 1 to 6+.

# **Examples**

 ${\tt StrelkaIDVCFFilesToCatalogAndPlotToPdf}$ 

Create ID (indel) catalog from Strelka ID VCF files and plot them to PDF

# Description

Create ID (indel) catalog from the Strelka ID VCFs specified by files and plot them to PDF

### Usage

```
StrelkaIDVCFFilesToCatalogAndPlotToPdf(files, ref.genome,
  region = "unknown", output.file)
```

### **Arguments**

files Character vector of file paths to the Strelka ID VCF files.

ref.genome A ref.genome argument as described in ICAMS.

region A character string designating a genomic region; see as.catalog and ICAMS.

output.file The name of the PDF file to be produced.

### **Details**

This function calls StrelkaIDVCFFilesToCatalog and PlotCatalogToPdf

### Value

An ID (indel) catalog and its graph plotted to PDF with specified file name. The ID (indel) catalog has attributes added. See as.catalog for more details.

#### Note

In ID (insertion and deletion) catalogs, deletion repeat sizes range from 0 to 5+, but for plotting and end-user documentation deletion repeat sizes range from 1 to 6+.

### **Examples**

StrelkaSBSVCFFilesToCatalog

Create SBS and DBS catalogs from Strelka SBS VCF files.

# **Description**

Create 3 SBS catalogs (96, 192, 1536) and 3 DBS catalogs (78, 136, 144) from the Strelka SBS VCFs specified by files

```
StrelkaSBSVCFFilesToCatalog(files, ref.genome, trans.ranges = NULL,
  region = "unknown")
```

### **Arguments**

files	Character vector	of file paths to i	the Strelka SBS	VCF files.

ref.genome A ref.genome argument as described in ICAMS.

trans.ranges a data.table which contains transcript range and strand information. Please

refer to TranscriptRanges for more details.

region A character string designating a genomic region; see as . catalog and ICAMS.

### **Details**

This function calls VCFsToSBSCatalogs and VCFsToDBSCatalogs.

### Value

A list of 3 SBS catalogs (one each for 96, 192, and 1536) and 3 DBS catalogs (one each for 78, 136, and 144). If trans.ranges = NULL, SBS 192 and DBS 144 catalog will not be generated. Each catalog has attributes added. See as .catalog for more details.

### Note

SBS 192 and DBS 144 catalog only contains mutations in transcribed regions.

# **Examples**

 ${\tt StrelkaSBSVCFFilesToCatalogAndPlotToPdf}$ 

Create SBS and DBS catalogs from Strelka SBS VCF files and plot them to PDF

# **Description**

Create 3 SBS catalogs (96, 192, 1536) and 3 DBS catalogs (78, 136, 144) from the Strelka SBS VCFs specified by files and plot them to PDF

```
StrelkaSBSVCFFilesToCatalogAndPlotToPdf(files, ref.genome,
    trans.ranges = NULL, region = "unknown", output.file)
```

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### **Arguments**

files Character vector of file paths to the Strelka SBS VCF files.

ref.genome A ref.genome argument as described in ICAMS.

trans.ranges a data.table which contains transcript range and strand information. Please

refer to TranscriptRanges for more details.

region A character string designating a genomic region; see as.catalog and ICAMS.

output.file The name of the PDF file to be produced.

#### **Details**

 $This \ function \ calls \ StrelkaSBSVCFFilesToCatalog \ and \ PlotCatalogToPdf$ 

### Value

A list of 3 SBS catalogs (one each for 96, 192, and 1536), 3 DBS catalogs (one each for 78, 136, and 144) and their graphs plotted to PDF with specified file name. If trans.ranges = NULL, SBS 192 and DBS 144 catalog will not be generated and plotted. Each catalog has attributes added. See as.catalog for more details.

### Note

SBS 192 and DBS 144 catalogs include only mutations in transcribed regions.

# **Examples**

TranscriptRanges

Transcript ranges data

# **Description**

Transcript ranges and strand information for a particular reference genome.

```
trans.ranges.GRCh37
trans.ranges.GRCh38
trans.ranges.GRCm38
```

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#### **Format**

A data.table which contains transcript range and strand information for a particular reference genome. colnames are chrom, start, end, strand, gene.name. It uses one-based coordinates.

### **Details**

This information is needed to generate catalogs that depend on transcriptional strand information, for example catalogs of class SBS192Catalog.

```
trans.ranges.GRCh37: Human GRCh37. trans.ranges.GRCh38: Human GRCh38. trans.ranges.GRCm38: Mouse GRCm38.
```

For these two tables, only genes that are associated with a CCDS ID are kept for transcriptional strand bias analysis.

```
This information is needed for StrelkaSBSVCFFilesToCatalog,
StrelkaSBSVCFFilesToCatalogAndPlotToPdf, MutectVCFFilesToCatalog,
MutectVCFFilesToCatalogAndPlotToPdf, VCFsToSBSCatalogs and VCFsToDBSCatalogs.
```

#### **Source**

```
ftp://ftp.ebi.ac.uk/pub/databases/gencode/Gencode_human/release_30/GRCh37_mapping/
gencode.v30lift37.annotation.gff3.gz
ftp://ftp.ebi.ac.uk/pub/databases/gencode/Gencode_human/release_30/gencode.v30.annotation.
gff3.gz
ftp://ftp.ebi.ac.uk/pub/databases/gencode/Gencode_mouse/release_M21/gencode.vM21.
annotation.gff3.gz
```

# **Examples**

```
trans.ranges.GRCh37
# chrom start end strand gene.name
    1 65419 71585
1 367640 368634
                 71585 +
                                OR4F5
    1
                           +
                                0R4F29
     1
        621059
                622053
                                OR4F16
       859308 879961
     1
                                SAMD11
    1 879583 894689
                                 NOC2L
           . . .
                    . . .
                          . . .
```

 ${\it TransformCatalog}$ 

Transform between counts and density spectrum catalogs and counts and density signature catalogs.

# **Description**

Transform between counts and density spectrum catalogs and counts and density signature catalogs.

```
TransformCatalog(catalog, target.ref.genome = NULL,
  target.region = NULL, target.catalog.type = NULL,
  target.abundance = NULL)
```

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### **Arguments**

catalog An SBS or DBS catalog as described in ICAMS; must **not** be an ID (indel) catalog.

target.ref.genome

A ref.genome argument as described in ICAMS. If NULL, then defaults to the ref.genome attribute of catalog.

target.region A region argument; see as.catalog and ICAMS. If NULL, then defaults to the region attribute of catalog.

target.catalog.type

A character string acting as a catalog type identifier, one of "counts", "density", "counts.signature", "density.signature"; see as.catalog. If NULL, then defaults to the catalog. type attribute of catalog.

target.abundance

A vector of counts different source K-mer sequences for mutations. See all.abundance. If NULL, then the function attempt to infer the target.abundace from the class of catalog and the values of the target.ref.genome, target.region, and target.catalog.type. It is an error if the inferred abundance is different from an non-NULL target.abundance.

### **Details**

Only the following transformations are legal:

- 1. counts -> counts (used to transform between the source abundance and target.abundance)
- 2. counts -> density
- 3. counts -> (counts.signature,density.signature)
- 4. density -> counts (the semantics are to infer the genome-wide or exome-wide counts based on the densities)
- 5. density -> density (a null operation, generates a warning)
- 6. density -> (counts.signature,density.signature)
- 7. counts.signature -> counts.signature (used to transform between the source abundance and target.abundance)
- 8. counts.signature -> density.signature
- 9. counts.signature -> (counts, density) (generates an error)
- 10. density.signature -> density.signature (a null operation, generates a warning)
- 11. density.signature -> counts.signature
- 12. density.signature -> (counts, density) (generates an error)

### Value

A catalog as defined in ICAMS.

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VCFsToDBSCatalogs

Create DBS catalogs from VCFs

# **Description**

Create a list of 3 catalogs (one each for DBS78, DBS144 and DBS136) out of the contents in list.of.DBS.vcfs. The VCFs must not contain any type of mutation other then DBSs.

### Usage

```
VCFsToDBSCatalogs(list.of.DBS.vcfs, ref.genome, trans.ranges = NULL,
  region = "unknown")
```

# **Arguments**

list.of.DBS.vcfs

List of in-memory data frames of pure DBS mutations – no SBS or 3+BS muta-

tions. The list names will be the sample ids in the output catalog.

ref.genome A ref.genome argument as described in ICAMS.

trans.ranges a data.table which contains transcript range and strand information. Please

refer to TranscriptRanges for more details.

region A character string designating a genomic region; see as . catalog and ICAMS.

### Value

A list of 3 DBS catalogs, one each for 78, 144, 136: catDBS78 catDBS144 catDBS136. If trans.ranges = NULL, DBS 144 catalog will not be generated. Each catalog has attributes added. See as.catalog for more details.

### Note

DBS 144 catalog only contains mutations in transcribed regions.

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VCFsToIDCatalogs

Create ID (insertion and deletion) catalog from ID VCFs

### **Description**

Create ID (insertion and deletion) catalog from ID VCFs

### Usage

```
VCFsToIDCatalogs(list.of.vcfs, ref.genome, region = "unknown")
```

# **Arguments**

list.of.vcfs List of in-memory VCFs. The list names will be the sample ids in the output

catalog.

ref.genome A ref.genome argument as described in ICAMS.

region A character string acting as a region identifier, one of "genome", "exome".

### Value

An S3 object containing an ID (indel) catalog with class "catalog". See as.catalog for more details.

### Note

In ID (insertion and deletion) catalogs, deletion repeat sizes range from 0 to 5+, but for plotting and end-user documentation deletion repeat sizes range from 1 to 6+.

### **Examples**

VCFsToSBSCatalogs

Create SBS catalogs from SBS VCFs

# Description

Create a list of 3 catalogs (one each for 96, 192, 1536) out of the contents in list.of.SBS.vcfs. The SBS VCFs must not contain DBSs, indels, or other types of mutations.

```
VCFsToSBSCatalogs(list.of.SBS.vcfs, ref.genome, trans.ranges = NULL,
  region = "unknown")
```

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### **Arguments**

list.of.SBS.vcfs

List of in-memory data frames of pure SBS mutations – no DBS or 3+BS muta-

tions. The list names will be the sample ids in the output catalog.

ref.genome A ref.genome argument as described in ICAMS.

trans.ranges a data.table which contains transcript range and strand information. Please

refer to TranscriptRanges for more details.

region A character string designating a genomic region; see as.catalog and ICAMS.

### Value

A list of 3 SBS catalogs, one each for 96, 192, 1536: catSBS96 catSBS192 catSBS1536. If trans.ranges = NULL, SBS 192 catalog will not be generated. Each catalog has attributes added. See as.catalog for more details.

#### Note

SBS 192 catalogs only contain mutations in transcribed regions.

# **Examples**

WriteCatalog

Write a catalog

### **Description**

Write a catalog to a file.

# Usage

```
WriteCatalog(catalog, file, strict = TRUE)
```

# **Arguments**

catalog A catalog as defined in ICAMS; see also as.catalog.

file The path to the file to be created.

strict If TRUE, do additional checks on the input, and stop if the checks fail.

# Details

See also ReadCatalog.

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### Note

In ID (insertion and deletion) catalogs, deletion repeat sizes range from 0 to 5+, but for plotting and end-user documentation deletion repeat sizes range from 1 to 6+.

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