

# Package ‘ICAMS’

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**Type** Package

**Title** In-depth Characterization and Analysis of Mutational Signatures ('ICAMS')

**Version** 2.1.1

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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.

**License** GPL-3

**URL** <https://github.com/steverozen/ICAMS>

**BugReports** <https://github.com/steverozen/ICAMS/issues>

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BSgenome,  
data.table,  
dplyr,  
GenomeInfoDb,  
GenomicRanges,  
graphics,  
grDevices,  
IRanges,  
RColorBrewer,  
stats,  
stringi,  
utils,  
zip

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BSgenome.Hsapiens.1000genomes.hs37d5,

BSgenome.Hsapiens.UCSC.hg38,

BSgenome.Mmusculus.UCSC.mm10

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all.abundance	<i>K-mer abundances.</i>
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### 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  
`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.
```

---

AnnotateDBSVCF	<i>Add sequence context and transcript information to an in-memory DBS VCF.</i>
----------------	---

---

### Description

Add sequence context and transcript information to an in-memory DBS VCF.

### Usage

```
AnnotateDBSVCF(DBS.vcf, ref.genome, trans.ranges = NULL)
```

## Arguments

<code>DBS.vcf</code>	An in-memory DBS VCF as a <code>data.frame</code> .
<code>ref.genome</code>	A <code>ref.genome</code> argument as described in <a href="#">ICAMS</a> .
<code>trans.ranges</code>	Optional. If <code>ref.genome</code> specifies one of the <a href="#">BSgenome</a> object <ol style="list-style-type: none"> <li>1. <a href="#">BSgenome.Hsapiens.1000genomes.hs37d5</a></li> <li>2. <a href="#">BSgenome.Hsapiens.UCSC.hg38</a></li> <li>3. <a href="#">BSgenome.Mmusculus.UCSC.mm10</a></li> </ol> then the function will infer <code>trans.ranges</code> automatically. Otherwise, user will need to provide the necessary <code>trans.ranges</code> . Please refer to <a href="#">TranscriptRanges</a> for more details. If <code>is.null(trans.ranges)</code> do not add transcript range information.

## Value

An in-memory DBS VCF as a `data.table`. This has been annotated with the sequence context (column name `seq.21bases`) and with transcript information in the form of a gene symbol (e.g. "TP53") and transcript strand. This information is in the columns `trans.start.pos`, `trans.end.pos`, `trans.strand`, `trans.Ensembl.gene.ID` and `trans.gene.symbol` in the output. These columns are not added if `is.null(trans.ranges)`.

## Examples

```
file <- c(system.file("extdata/Strelka-SBS-vcf",
                     "Strelka.SBS.GRCh37.vcf",
                     package = "ICAMS"))
list.of.vcfs <- ReadAndSplitStrelkaSBSVCFs(file)
DBS.vcf <- list.of.vcfs$DBS.vcfs[[1]]
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  annotated.DBVCF <- AnnotateDBSVCF(DBS.vcf, ref.genome = "hg19",
                                   trans.ranges = trans.ranges.GRCh37)}
```

---

AnnotateIDVCF	<i>Add sequence context to an in-memory ID (insertion/deletion) VCF, and confirm that they match the given reference genome.</i>
---------------	--

---

## Description

Add sequence context to an in-memory ID (insertion/deletion) VCF, and confirm that they match the given reference genome.

## Usage

```
AnnotateIDVCF(ID.vcf, ref.genome, flag.mismatches = 0)
```

**Arguments**

<code>ID.vcf</code>	An in-memory ID (insertion/deletion) VCF as a <code>data.frame</code> . This function expects that there is a "context base" to the left, for example <code>REF = ACG</code> , <code>ALT = A</code> (deletion of CG) or <code>REF = A</code> , <code>ALT = ACC</code> (insertion of CC).
<code>ref.genome</code>	A <code>ref.genome</code> argument as described in <a href="#">ICAMS</a> .
<code>flag.mismatches</code>	If $> 0$ , then if there are mismatches to references, generate messages showing the mismatched rows and continue. Otherwise stop if there are mismatched rows.

**Value**

A data frame with 2 new columns added to the input data frame:

1. `seq.context` The sequence embedding the variant.
2. `seq.context.width` The width of `seq.context` to the left

**Examples**

```
file <- c(system.file("extdata/Strelka-ID-vcf/",
                     "Strelka.ID.GRCh37.vcf",
                     package = "ICAMS"))
ID.vcf <- ReadStrelkaIDVCFs(file)[[1]]
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  annotated.ID.vcf <- AnnotateIDVCF(ID.vcf, ref.genome = "hg19")
}
```

---

AnnotateSBSVCF	<i>Add sequence context and transcript information to an in-memory SBS VCF.</i>
----------------	---

---

**Description**

Add sequence context and transcript information to an in-memory SBS VCF.

**Usage**

```
AnnotateSBSVCF(SBS.vcf, ref.genome, trans.ranges = NULL)
```

**Arguments**

<code>SBS.vcf</code>	An in-memory SBS VCF as a <code>data.frame</code> .
<code>ref.genome</code>	A <code>ref.genome</code> argument as described in <a href="#">ICAMS</a> .
<code>trans.ranges</code>	Optional. If <code>ref.genome</code> specifies one of the <a href="#">BSgenome</a> object <ol style="list-style-type: none"> <li>1. <a href="#">BSgenome.Hsapiens.1000genomes.hs37d5</a></li> <li>2. <a href="#">BSgenome.Hsapiens.UCSC.hg38</a></li> <li>3. <a href="#">BSgenome.Mmusculus.UCSC.mm10</a></li> </ol> then the function will infer <code>trans.ranges</code> automatically. Otherwise, user will need to provide the necessary <code>trans.ranges</code> . Please refer to <a href="#">TranscriptRanges</a> for more details. If <code>is.null(trans.ranges)</code> do not add transcript range information.

**Value**

An in-memory SBS VCF as a `data.table`. This has been annotated with the sequence context (column name `seq.21bases`) and with transcript information in the form of a gene symbol (e.g. "TP53") and transcript strand. This information is in the columns `trans.start.pos`, `trans.end.pos`, `trans.strand`, `trans.Ensembl.gene.ID` and `trans.gene.symbol` in the output. These columns are not added if `is.null(trans.ranges)`.

**Examples**

```
file <- c(system.file("extdata/Strelka-SBS-vcf",
                     "Strelka.SBS.GRCh37.vcf",
                     package = "ICAMS"))
list.of.vcfs <- ReadAndSplitStrelkaSBSVCFs(file)
SBS.vcf <- list.of.vcfs$SBS.vcfs[[1]]
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  annotated.SBS.vcf <- AnnotateSBSVCF(SBS.vcf, ref.genome = "hg19",
                                     trans.ranges = trans.ranges.GRCh37)}
```

as.catalog

*Create a catalog from a matrix, data.frame, or vector.***Description**

Create a catalog from a matrix, data.frame, or vector.

**Usage**

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

**Arguments**

- |              |  |
|--------------|--|
| object       | A numeric matrix, numeric data.frame, or vector. If a vector, converted to a 1-column matrix with rownames taken from the element names of the vector and with column name "Unknown". If argument <code>infer.rownames</code> is FALSE than this argument must have rownames to denote the mutation types. See <a href="#">CatalogRowOrder</a> for more details. |
| ref.genome   | A ref.genome argument as described in <a href="#">ICAMS</a> .  |
| region       | A character string designating a region, one of genome, transcript, exome, unknown; see <a href="#">ICAMS</a> .  |
| 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 <a href="#">ICAMS</a> . The argument abundance should contain the counts of different source sequences for mutations in the same format as the numeric vectors in <a href="#">all.abundance</a> .   |

`infer.rownames` If TRUE, and object has no rownames, then assume the rows of object are in the correct order and add the rownames implied by the number of rows in object (e.g. rownames for SBS 192 if there are 192 rows). If TRUE, **be sure the order of rows is correct.**

### Value

A catalog as described in [ICAMS](#).

### Examples

```
# Create an SBS96 catalog with all mutation counts equal to 1.
object <- matrix(1, nrow = 96, ncol = 1,
                 dimnames = list(catalog.row.order$SBS96))
catSBS96 <- as.catalog(object)
```

---

Canonicalize1Del	<i>Given a deletion and its sequence context, categorize it.</i>
------------------	--

---

### Description

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

### Usage

```
Canonicalize1Del(context, del.seq, pos, trace = 0)
```

### Arguments

<code>context</code>	The deleted sequence plus ample surrounding sequence on each side (at least as long as <code>del.seq</code> ).
<code>del.seq</code>	The deleted sequence in context.
<code>pos</code>	The position of <code>del.seq</code> in context.
<code>trace</code>	If > 0, then generate messages tracing how the computation is carried out.

### Details

See [https://github.com/steverozen/ICAMS/raw/master/data-raw/PCAWG7\\_indel\\_classification\\_2017\\_12\\_08.xlsx](https://github.com/steverozen/ICAMS/raw/master/data-raw/PCAWG7_indel_classification_2017_12_08.xlsx) for additional information on deletion mutation classification.

This function first handles deletions in homopolymers, then handles deletions in simple repeats with longer repeat units, (e.g. CACACACA, see [FindMaxRepeatDel](#)), and if the deletion is not in a simple repeat, looks for microhomology (see [FindDeIMH](#)).

See the code for unexported function [CanonicalizeID](#) and the functions it calls for handling of insertions.

### Value

A string that is the canonical representation of the given deletion type. Return NA and raise a warning if there is an un-normalized representation of the deletion of a repeat unit. See [FindDeIMH](#) for details. (This seems to be very rare.)

## Examples

```
Canonicalize1Del("xyAAAqr", del.seq = "A", pos = 3) # "DEL:T:1:2"
Canonicalize1Del("xyAAAqr", del.seq = "A", pos = 4) # "DEL:T:1:2"
Canonicalize1Del("xyAqr", del.seq = "A", pos = 3)   # "DEL:T:1:0"
```

---

CatalogRowOrder	<i>Standard order of row names in a catalog.</i>
-----------------	--

---

## 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
catalog.row.order.sp
```

## Format

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

An object of class list of length 8.

An object of class list of length 1.

## Note

In ID (small 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+. In ID83 catalogs, deletion repeat sizes range from 0 to 5.

## Examples

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

catalog.row.order.sp$ID83
# "DEL:C:1:0" "DEL:C:1:1" "DEL:C:1:2" "DEL:C:1:3" ...
# There are altogether 83 row names to denote the mutation types
# in ID83 catalog.
```



---

CollapseCatalog	<i>"Collapse" a catalog.</i>
-----------------	------------------------------

---

### 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.

### Usage

```
Collapse192CatalogTo96(catalog)
```

```
Collapse1536CatalogTo96(catalog)
```

```
Collapse144CatalogTo78(catalog)
```

### Arguments

catalog      A catalog as defined in [ICAMS](#).

### Value

A catalog as defined in [ICAMS](#).

### Examples

```
# Create an SBS192 catalog and collapse it to an SBS96 catalog
object <- matrix(1, nrow = 192, ncol = 1,
                 dimnames = list(catalog.row.order$SBS192))
catSBS192 <- as.catalog(object, region = "transcript")
catSBS96 <- Collapse192CatalogTo96(catSBS192)
```

---

FindDelMH	<i>Return the length of microhomology at a deletion.</i>
-----------	--

---

### Description

Return the length of microhomology at a deletion.

### Usage

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

**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, then generate various messages showing how the computation is carried out.
warn.cryptic	if TRUE generating a warning if there is a cryptic repeat (see the example).

**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:

```

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

```

Presumed repair mechanism leading to this:

```

....
GGCTAGAACTAGTT
CCGATCTTGATCAA

```

=>

```

....
GGCTAG      TT
CC      GATCAA
      ....

```

=>

```

GGCTAGTT
CCGATCAA

```

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

```

GGC-----TAGTT  GGCTAGTT  GGC[TAGAAC]TAGTT
                        * --- * ---

```

```

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

```

```

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

```

```

          *** - *** -
GGCTAG-----TT GGCTAGTT GGCTAG[AACTAG]TT
          *****

```

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).

### Warning

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

```

GACTAGCTAGTT
GACTA----GTT GACTAGTT GACTA[GCTA]GTT
          *** -*** -

```

is really a repeat

```

GACTAG----TT GACTAGTT GACTAG[CTAG]TT
          ***** ----
GACT----AGTT GACTAGTT GACT[AGCT]AGTT
          ** ----* --

```

**This function only flags these "cryptic repeats" with a -1 return; it does not figure out the repeat extent.**

### Value

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

### Examples

```

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

# A cryptic repeat
#
# TAAATTATTTATTAATTTATTG
# TAAATTA----TTAATTTATTG = TAAATTATTAATTTATTG
#
# equivalent to
#
# TAAATTATTTATTAATTTATTG
# TAAAT----TATTAATTTATTG = TAAATTATTAATTTATTG
#

```

```
# and
#
# TAAATTATTTATTAATTTATTG
# TAAA----TTATTAATTTATTG = TAAATTATTAATTTATTG

FindDelMH("TAAATTATTTATTAATTTATTG", "TTTA", 8, warn.cryptic = FALSE) # -1
```

---

FindMaxRepeatDel	<i>Return the number of repeat units in which a deletion is embedded.</i>
------------------	---

---

## Description

Return the number of repeat units in which a deletion is embedded.

## Usage

```
FindMaxRepeatDel(context, rep.unit.seq, pos)
```

## Arguments

context	A string that embeds rep.unit.seq at position pos
rep.unit.seq	A substring of context at pos to pos + nchar(rep.unit.seq) - 1, which is the repeat unit sequence.
pos	The position of rep.unit.seq in context.

## Details

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

For example `FindMaxRepeatDel("xyaczt", "ac", 3)` returns 0.

If `substr(context, pos, pos + nchar(rep.unit.seq) - 1) != rep.unit.seq` then stop.

If this function returns 0, then it is necessary to look for microhomology using the function [FindDelMH](#).

### Warning

This function depends on the variant caller having "aligned" the deletion within the context of the repeat.

For example, a deletion of CAG in the repeat

```
GTCAGCAGCATGT
```

can have 3 "aligned" representations as follows:

```
CT---CAGCAGGT
CTCAG---CAGGT
CTCAGCAG---GT
```

In these cases this function will return 2. (Please note that the return value does not include the rep.unit.seq in the count.)

However, the same deletion can also have an "unaligned" representation, such as

```
CTCAGC---AGGT
```

(a deletion of AGC).

In this case this function will return 1 (a deletion of AGC in a 2-element repeat of AGC).

**Value**

The number of repeat units in which `rep.unit.seq` is embedded, not including the input `rep.unit.seq` in the count.

**Examples**

```
FindMaxRepeatDel("xyACACzt", "AC", 3) # 1
FindMaxRepeatDel("xyACACzt", "CA", 4) # 0
```

---

GeneExpressionData	<i>Example gene expression data from two cell lines.</i>
--------------------	--

---

**Description**

This data is designed to be used as an example in function [PlotTransBiasGeneExp](#) and [PlotTransBiasGeneExpToPdf](#).

**Usage**

```
gene.expression.data.HepG2

gene.expression.data.MCF10A
```

**Format**

A [data.table](#) which contains the expression values of genes.

An object of class `data.table` (inherits from `data.frame`) with 57736 rows and 4 columns.

An object of class `data.table` (inherits from `data.frame`) with 57736 rows and 4 columns.

**Examples**

```
gene.expression.data.HepG2
# Ensembl.gene.ID gene.symbol counts      TPM
# ENSG000000000003      TSPAN6    6007 33.922648455
# ENSG000000000005      TNMD       0  0.000000000
# ENSG000000000419      DPM1    4441 61.669371091
# ENSG000000000457      SCYL3    1368  3.334619195
# ENSG000000000460    C1orf112    916  2.416263423
#                ...      ...      ...      ...
```

---

GetVAF	<i>Extract the VAFs (variant allele frequencies) and read depth information from a VCF file.</i>
--------	--

---

### Description

Extract the VAFs (variant allele frequencies) and read depth information from a VCF file.

### Usage

```
GetStrelkaVAF(vcf, name.of.VCF = NULL)
```

```
GetMutectVAF(vcf, name.of.VCF = NULL, tumor.col.name = NA)
```

### Arguments

vcf	Said VCF as a data.frame.
name.of.VCF	Name of the VCF file.
tumor.col.name	Optional. Name of the column in VCF which contains the tumor sample information. It <b>must</b> have quotation marks. If tumor.col.name is equal to NA(default), this function will use the 10th column to calculate VAFs.

### Value

The original vcf with two additional columns added which contain the VAF(variant allele frequency) and read depth information.

### Examples

```
file <- c(system.file("extdata/Strelka-SBS-vcf",
                     "Strelka.SBS.GRCh37.vcf",
                     package = "ICAMS"))
MakeDataFrameFromStrelkaSBSVCF <-
  getFromNamespace("MakeDataFrameFromStrelkaSBSVCF", "ICAMS")
df <- MakeDataFrameFromStrelkaSBSVCF(file)
df1 <- GetStrelkaVAF(df)
```

---

ICAMS	<i>ICAMS: In-depth Characterization and Analysis of Mutational Signatures</i>
-------	---

---

### 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 S3 objects of class `matrix` and one of several additional classes that specify the types of the mutations represented in the catalog. The possible additional class is one of

- `SBS96Catalog` (strand-agnostic single base substitutions in trinucleotide context)
- `SBS192Catalog` (transcription-stranded single-base substitutions in trinucleotide context)
- `SBS1536Catalog`
- `DBS78Catalog`
- `DBS144Catalog`
- `DBS136Catalog`
- `IndelCatalog`

`as.catalog` is the main constructor.

Conceptually, a catalog also has one of the following types, indicated by 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
  - 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 [TransformCatalog](#) for more information. Abundances logically depend on the species in question and on the part of the genome being analyzed.

In "ICAMS" abundances can sometimes be inferred from the `catalog` class attribute and the function arguments `region`, `ref.genome`, and `catalog.type`. Otherwise abundances 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 an ID (small insertion and deletion) catalog from the Strelka ID VCFs.
3. [MutectVCFFilesToCatalog](#) creates 3 SBS catalogs (96, 192, 1536), 3 DBS catalogs (78, 136, 144) and ID (small insertion and deletion) 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 (small insertion and deletion) 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.

### Wrapper functions to create a zip file which contains catalogs and plot PDFs from VCF files

1. [StrelkaSBSVCFFilesToZipFile](#) creates a zip file which contains SBS and DBS catalogs and plot PDFs from Strelka SBS VCF files.
2. [StrelkaIDVCFFilesToZipFile](#) creates a zip file which contains ID (small insertion and deletion) catalog and plot PDF from Strelka ID VCF files.
3. [MutectVCFFilesToZipFile](#) creates a zip file which contains SBS, DBS and ID catalogs and plot PDFs from Mutect VCF files.

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

Many functions take the argument `ref.genome`.

To create a mutational spectrum catalog from a VCF file, ICAMS needs the reference genome sequence that matches the VCF file. The `ref.genome` argument provides this.

`ref.genome` must be one of

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", which specify [BSgenome.Hsapiens.UCSC.hg38](#).
3. The strings "hg19" or "GRCh37", which specify [BSgenome.Hsapiens.1000genomes.hs37d5](#).
4. The strings "mm10" or "GRCm38", which specify [BSgenome.Mmusculus.UCSC.mm10](#).

All needed reference genomes must be installed separately by the user. Further instructions are at <https://bioconductor.org/packages/release/bioc/html/BSgenome.html>.



Use of ICAMS with reference genomes other than the 2 human genomes and 1 mouse genome specified above is restricted to `catalog.type` of `counts` or `counts.signature` unless the user also creates the necessary abundance vectors. See [all.abundance](#).

Use [available.genomes\(\)](#) to get the list of available genomes.

### 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 fine-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. 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.

3. [GeneExpressionData](#) Example gene expression data from two cell lines.

---

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",
  names.of.VCFs = NULL,
  tumor.col.names = NA,
  flag.mismatches = 0
)
```

## Arguments

<code>files</code>	Character vector of file paths to the Mutect VCF files.
<code>ref.genome</code>	A <code>ref.genome</code> argument as described in <a href="#">ICAMS</a> .
<code>trans.ranges</code>	Optional. If <code>ref.genome</code> specifies one of the <a href="#">BSgenome</a> object <ol style="list-style-type: none"> <li>1. <a href="#">BSgenome.Hsapiens.1000genomes.hs37d5</a></li> <li>2. <a href="#">BSgenome.Hsapiens.UCSC.hg38</a></li> <li>3. <a href="#">BSgenome.Mmusculus.UCSC.mm10</a></li> </ol> then the function will infer <code>trans.ranges</code> automatically. Otherwise, user will need to provide the necessary <code>trans.ranges</code> . Please refer to <a href="#">TranscriptRanges</a> for more details. If <code>is.null(trans.ranges)</code> do not add transcript range information.
<code>region</code>	A character string designating a genomic region; see <a href="#">as.catalog</a> and <a href="#">ICAMS</a> .
<code>names.of.VCFs</code>	Optional. Character vector of names of the VCF files. The order of names in <code>names.of.VCFs</code> should match the order of VCF file paths in <code>files</code> . If <code>NULL</code> (default), this function will remove all of the path up to and including the last path separator (if any) in <code>files</code> and file paths without extensions (and the leading dot) will be used as the names of the VCF files.
<code>tumor.col.names</code>	Optional. Character vector of column names in VCFs which contain the tumor sample information. The order of names in <code>tumor.col.names</code> should match the order of VCFs specified in <code>files</code> . If <code>tumor.col.names</code> is equal to <code>NA</code> (default), this function will use the 10th column in all the VCFs to calculate VAFs. See <a href="#">GetMutectVAF</a> for more details.

`flag.mismatches`

Optional. If  $> 0$ , then if there are mismatches to references in the ID (insertion/deletion) VCF, generate messages showing the mismatched rows and continue. Otherwise stop if there are mismatched rows. See [AnnotateIDVCF](#) for more details.

## 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.

## Comments

To add or change attributes of the catalog, you can use function [attr](#). For example, `attr(catalog, "abundance") <- custom.abundance`.

## Note

SBS 192 and DBS 144 catalogs include only mutations in transcribed regions. In ID (small 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

```
file <- c(system.file("extdata/Mutect-vcf",
                     "Mutect.GRCh37.vcf",
                     package = "ICAMS"))
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  catalogs <- MutectVCFFilesToCatalog(file, ref.genome = "hg19",
                                     trans.ranges = trans.ranges.GRCh37,
                                     region = "genome")}
```

---

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",
  names.of.VCFs = NULL,
  tumor.col.names = NA,
  output.file = "",
  flag.mismatches = 0
)
```

## Arguments

<code>files</code>	Character vector of file paths to the Mutect VCF files.
<code>ref.genome</code>	A <code>ref.genome</code> argument as described in <a href="#">ICAMS</a> .
<code>trans.ranges</code>	Optional. If <code>ref.genome</code> specifies one of the <a href="#">BSgenome</a> object <ol style="list-style-type: none"> <li>1. <a href="#">BSgenome.Hsapiens.1000genomes.hs37d5</a></li> <li>2. <a href="#">BSgenome.Hsapiens.UCSC.hg38</a></li> <li>3. <a href="#">BSgenome.Mmusculus.UCSC.mm10</a></li> </ol> then the function will infer <code>trans.ranges</code> automatically. Otherwise, user will need to provide the necessary <code>trans.ranges</code> . Please refer to <a href="#">TranscriptRanges</a> for more details. If <code>is.null(trans.ranges)</code> do not add transcript range information.
<code>region</code>	A character string designating a genomic region; see <a href="#">as.catalog</a> and <a href="#">ICAMS</a> .
<code>names.of.VCFs</code>	Optional. Character vector of names of the VCF files. The order of names in <code>names.of.VCFs</code> should match the order of VCF file paths in <code>files</code> . If <code>NULL</code> (default), this function will remove all of the path up to and including the last path separator (if any) in <code>files</code> and file paths without extensions (and the leading dot) will be used as the names of the VCF files.
<code>tumor.col.names</code>	Optional. Character vector of column names in VCFs which contain the tumor sample information. The order of names in <code>tumor.col.names</code> should match the order of VCFs specified in <code>files</code> . If <code>tumor.col.names</code> is equal to <code>NA</code> (default), this function will use the 10th column in all the VCFs to calculate VAFs. See <a href="#">GetMutectVAF</a> for more details.
<code>output.file</code>	Optional. The base name of the PDF files to be produced; multiple files will be generated, each ending in <code>x.pdf</code> , where <code>x</code> indicates the type of catalog plotted in the file.
<code>flag.mismatches</code>	Optional. If <code>&gt; 0</code> , then if there are mismatches to references in the ID (insertion/deletion) VCF, generate messages showing the mismatched rows and continue. Otherwise stop if there are mismatched rows. See <a href="#">AnnotateIDVCF</a> for more details.

## 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.

**Comments**

To add or change attributes of the catalog, you can use function [attr](#).  
For example, `attr(catalog, "abundance") <- custom.abundance`.

**Note**

SBS 192 and DBS 144 catalogs include only mutations in transcribed regions. In ID (small 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**

```
file <- c(system.file("extdata/Mutect-vcf",
                     "Mutect.GRCh37.vcf",
                     package = "ICAMS"))
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  catalogs <-
    MutectVCFFilesToCatalogAndPlotToPdf(file, ref.genome = "hg19",
                                         trans.ranges = trans.ranges.GRCh37,
                                         region = "genome",
                                         output.file =
                                           file.path(tempdir(), "Mutect"))}
```

---

MutectVCFFilesToZipFile

*Create a zip file which contains catalogs and plot PDFs 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 `dir`, save the catalogs as CSV files, plot them to PDF and generate a zip archive of all the output files.

**Usage**

```
MutectVCFFilesToZipFile(
  dir,
  zipfile,
  ref.genome,
  trans.ranges = NULL,
  region = "unknown",
  names.of.VCFs = NULL,
  tumor.col.names = NA,
  base.filename = "",
```

```
    flag.mismatches = 0
  )
```

### Arguments

<code>dir</code>	Pathname of the directory which contains the Mutect VCF files. Each Mutect VCF <b>must</b> have a file extension ".vcf" (case insensitive) and share the <b>same</b> <code>ref.genome</code> and <code>region</code> .
<code>zipfile</code>	Pathname of the zip file to be created.
<code>ref.genome</code>	A <code>ref.genome</code> argument as described in <a href="#">ICAMS</a> .
<code>trans.ranges</code>	Optional. If <code>ref.genome</code> specifies one of the <a href="#">BSgenome</a> object <ol style="list-style-type: none"> <li>1. <a href="#">BSgenome.Hsapiens.1000genomes.hs37d5</a></li> <li>2. <a href="#">BSgenome.Hsapiens.UCSC.hg38</a></li> <li>3. <a href="#">BSgenome.Mmusculus.UCSC.mm10</a></li> </ol> then the function will infer <code>trans.ranges</code> automatically. Otherwise, user will need to provide the necessary <code>trans.ranges</code> . Please refer to <a href="#">TranscriptRanges</a> for more details. If <code>is.null(trans.ranges)</code> do not add transcript range information.
<code>region</code>	A character string designating a genomic region; see <a href="#">as.catalog</a> and <a href="#">ICAMS</a> .
<code>names.of.VCFs</code>	Optional. Character vector of names of the VCF files. The order of names in <code>names.of.VCFs</code> should match the order of VCFs listed in <code>dir</code> . If <code>NULL</code> (default), this function will remove all of the path up to and including the last path separator (if any) in <code>dir</code> and file paths without extensions (and the leading dot) will be used as the names of the VCF files.
<code>tumor.col.names</code>	Optional. Character vector of column names in VCFs which contain the tumor sample information. The order of names in <code>tumor.col.names</code> should match the order of VCFs listed in <code>dir</code> . If <code>tumor.col.names</code> is equal to <code>NA</code> (default), this function will use the 10th column in all the VCFs to calculate VAFs. See <a href="#">GetMutectVAF</a> for more details.
<code>base.filename</code>	Optional. The base name of the CSV and PDF files to be produced; multiple files will be generated, each ending in <code>x.csv</code> or <code>x.pdf</code> , where <code>x</code> indicates the type of catalog.
<code>flag.mismatches</code>	Optional. If <code>&gt; 0</code> , then if there are mismatches to references in the ID (insertion/deletion) VCF, generate messages showing the mismatched rows and continue. Otherwise stop if there are mismatched rows. See <a href="#">AnnotateIDVCF</a> for more details.

### Details

This function calls [MutectVCFFilesToCatalog](#), [PlotCatalogToPdf](#), [WriteCatalog](#) and [zipr](#).

### 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 Indel catalog. 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.

## Comments

To add or change attributes of the catalog, you can use function [attr](#).  
For example, `attr(catalog, "abundance") <- custom.abundance`.

## Note

SBS 192 and DBS 144 catalogs include only mutations in transcribed regions. In ID (small 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

```
dir <- c(system.file("extdata/Mutect-vcf",
                    package = "ICAMS"))
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  catalogs <-
    MutectVCFFilesToZipFile(dir,
                           zipfile = file.path(tempdir(), "test.zip"),
                           ref.genome = "hg19",
                           trans.ranges = trans.ranges.GRCh37,
                           region = "genome",
                           base.filename = "Mutect")
  unlink(file.path(tempdir(), "test.zip"))
}
```

---

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,
  ylim = NULL
)
```

## Arguments

catalog	A catalog as defined in <a href="#">ICAMS</a> with attributes added. See <a href="#">as.catalog</a> 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	Has the usual meaning. Taken from <code>par("cex")</code> by default. Only implemented for <b>SBS96Catalog</b> , <b>SBS192Catalog</b> and <b>DBS144Catalog</b> .
grid	A logical value indicating whether to draw grid lines. Only implemented for <b>SBS96Catalog</b> .
upper	A logical value indicating whether to draw horizontal lines and the names of major mutation class on top of graph. Only implemented for <b>SBS96Catalog</b> .
xlabels	A logical value indicating whether to draw x axis labels. Only implemented for <b>SBS96Catalog</b> . If FALSE then plot x axis tick marks; set <code>par(tck = 0)</code> to suppress.
ylim	Has the usual meaning. Only implemented for <b>SBS96Catalog</b> and <b>IndelCatalog</b> .

### Value

A list whose first element is a logic value indicating whether the plot is successful. For **SBS96Catalog**, the list will have a second element, which is a numeric vector giving the coordinates of all the bar midpoints drawn, useful for adding to the graph. For **SBS192Catalog** with "counts" catalog.type and non-NULL abundance, the list will have a second element which is a list containing the strand bias statistics.

### Comments

For **SBS192Catalog** with "counts" catalog.type and non-NULL abundance, the strand bias statistics are Benjamini-Hochberg q-values based on two-sided binomial tests of the mutation counts on the transcribed and untranscribed strands relative to the actual abundances of C and T on the transcribed strand. On the SBS12 plot, asterisks indicate q-values as follows \*,  $Q < 0.05$ ; \*\*,  $Q < 0.01$ ; \*\*\*,  $Q < 0.001$ .

### 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

```
file <- system.file("extdata",
                    "strelka.regress.cat.sbs.96.csv",
                    package = "ICAMS")
catSBS96 <- ReadCatalog(file)
colnames(catSBS96) <- "sample"
PlotCatalog(catSBS96)
```

---

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,
  ylim = NULL
)
```

**Arguments**

catalog	A catalog as defined in <a href="#">ICAMS</a> with attributes added. See <a href="#">as.catalog</a> for more details.
file	The name of the PDF file to be produced.
plot.SBS12	Only meaningful for class <code>SBS192Catalog</code> ; 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	Has the usual meaning. A default value has been used by the program internally. Only implemented for <code>SBS96Catalog</code> , <code>SBS192Catalog</code> and <code>DBS144Catalog</code> .
grid	A logical value indicating whether to draw grid lines. Only implemented for <code>SBS96Catalog</code> .
upper	A logical value indicating whether to draw horizontal lines and the names of major mutation class on top of graph. Only implemented for <code>SBS96Catalog</code> .
xlabels	A logical value indicating whether to draw x axis labels. Only implemented for <code>SBS96Catalog</code> . If FALSE then plot x axis tick marks; set <code>par(tck = 0)</code> to suppress.
ylim	Has the usual meaning. Only implemented for <code>SBS96Catalog</code> and <code>IndelCatalog</code> .

**Value**

A list whose first element is a logic value indicating whether the plot is successful. For **SBS192Catalog** with "counts" catalog.type and non-null abundance, the list will have a second element which is a list containing the strand bias statistics.

**Comments**

For **SBS192Catalog** with "counts" catalog.type and non-NULL abundance, the strand bias statistics are Benjamini-Hochberg q-values based on two-sided binomial tests of the mutation counts on the transcribed and untranscribed strands relative to the actual abundances of C and T on the transcribed strand. On the SBS12 plot, asterisks indicate q-values as follows \*,  $Q < 0.05$ ; \*\*,  $Q < 0.01$ ; \*\*\*,  $Q < 0.001$ .

**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

```
file <- system.file("extdata",
                    "strelka.regress.cat.sbs.96.csv",
                    package = "ICAMS")
catSBS96 <- ReadCatalog(file)
colnames(catSBS96) <- "sample"
PlotCatalogToPdf(catSBS96, file = file.path(tempdir(), "test.pdf"))
```

---

PlotTransBiasGeneExp    *Plot transcription strand bias with respect to gene expression values.*

---

## Description

Plot transcription strand bias with respect to gene expression values.

## Usage

```
PlotTransBiasGeneExp(
  annotated.SBS.vcf,
  expression.data,
  Ensembl.gene.ID.col,
  expression.value.col,
  num.of.bins,
  plot.type,
  damaged.base = NULL,
  ymax = NULL
)
```

## Arguments

annotated.SBS.vcf	An SBS VCF annotated by <a href="#">AnnotateSBSVCF</a> . It <b>must</b> have transcript range information added.
expression.data	A <a href="#">data.table</a> which contains the expression values of genes. See <a href="#">GeneExpressionData</a> for more details.
Ensembl.gene.ID.col	Name of column which has the Ensembl gene ID information in expression.data.
expression.value.col	Name of column which has the gene expression values in expression.data.
num.of.bins	The number of bins that will be plotted on the graph.
plot.type	A character string indicating one mutation type to be plotted. It should be one of "C>A", "C>G", "C>T", "T>A", "T>C", "T>G".
damaged.base	One of NULL, "purine" or "pyrimidine". This function allocates approximately equal numbers of mutations from damaged.base into each of num.of.bins bin by expression level. E.g. if damaged.base is "purine", then mutations from A and G will be allocated in approximately equal numbers to each expression-level bin. The rationale for the name damaged.base is that the direction of strand bias is a result of whether the damage occurs on a purine or pyrimidine. If NULL, the function attempts to infer the damaged.base based on mutation counts.

**y**max                      Limit for the y axis. If not specified, it defaults to NULL and the y axis limit equals 1.5 times of the maximum mutation counts in a specific mutation type.

### Value

A list whose first element is a logic value indicating whether the plot is successful. The second element is a named numeric vector containing the p-values printed on the plot.

### Note

The p-values are calculated by logistic regression using function `glm`. The dependent variable is labeled "1" and "0" if the mutation from annotated.SBS.vcf falls onto the untranscribed and transcribed strand respectively. The independent variable is the binary logarithm of the gene expression value from expression.data plus one, i.e.  $\log_2(x + 1)$  where  $x$  stands for gene expression value.

### Examples

```
file <- c(system.file("extdata/Strelka-SBS-vcf/",
                     "Strelka.SBS.GRCh37.vcf",
                     package = "ICAMS"))
list.of.vcfs <- ReadAndSplitStrelkaSBSVCFs(file)
SBS.vcf <- list.of.vcfs$SBS.vcfs[[1]]
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  annotated.SBS.vcf <- AnnotateSBSVCF(SBS.vcf, ref.genome = "hg19",
                                     trans.ranges = trans.ranges.GRCh37)
  PlotTransBiasGeneExp(annotated.SBS.vcf = annotated.SBS.vcf,
                       expression.data = gene.expression.data.HepG2,
                       Ensembl.gene.ID.col = "Ensembl.gene.ID",
                       expression.value.col = "TPM",
                       num.of.bins = 4, plot.type = "C>A")
}
```

---

PlotTransBiasGeneExpToPdf

*Plot transcription strand bias with respect to gene expression values to a PDF file.*

---

### Description

Plot transcription strand bias with respect to gene expression values to a PDF file.

### Usage

```
PlotTransBiasGeneExpToPdf(
  annotated.SBS.vcf,
  file,
  expression.data,
  Ensembl.gene.ID.col,
  expression.value.col,
  num.of.bins,
  plot.type = c("C>A", "C>G", "C>T", "T>A", "T>C", "T>G"),
  damaged.base = NULL
)
```

## Arguments

<code>annotated.SBS.vcf</code>	An SBS VCF annotated by <a href="#">AnnotateSBSVCF</a> . It <b>must</b> have transcript range information added.
<code>file</code>	The name of output file.
<code>expression.data</code>	A <a href="#">data.table</a> which contains the expression values of genes. See <a href="#">GeneExpressionData</a> for more details.
<code>Ensembl.gene.ID.col</code>	Name of column which has the Ensembl gene ID information in <code>expression.data</code> .
<code>expression.value.col</code>	Name of column which has the gene expression values in <code>expression.data</code> .
<code>num.of.bins</code>	The number of bins that will be plotted on the graph.
<code>plot.type</code>	A vector of character indicating types to be plotted. It can be one or more types from "C>A", "C>G", "C>T", "T>A", "T>C", "T>G". The default is to print all the six mutation types.
<code>damaged.base</code>	One of NULL, "purine" or "pyrimidine". This function allocates approximately equal numbers of mutations from <code>damaged.base</code> into each of <code>num.of.bins</code> bin by expression level. E.g. if <code>damaged.base</code> is "purine", then mutations from A and G will be allocated in approximately equal numbers to each expression-level bin. The rationale for the name <code>damaged.base</code> is that the direction of strand bias is a result of whether the damage occurs on a purine or pyrimidine. If NULL, the function attempts to infer the <code>damaged.base</code> based on mutation counts.

## Value

A list whose first element is a logic value indicating whether the plot is successful. The second element is a named numeric vector containing the p-values printed on the plot.

## Note

The p-values are calculated by logistic regression using function [glm](#). The dependent variable is labeled "1" and "0" if the mutation from `annotated.SBS.vcf` falls onto the untranscribed and transcribed strand respectively. The independent variable is the binary logarithm of the gene expression value from `expression.data` plus one, i.e.  $\log_2(x + 1)$  where  $x$  stands for gene expression value.

## Examples

```
file <- c(system.file("extdata/Strelka-SBS-vcf/",
                     "Strelka.SBS.GRCh37.vcf",
                     package = "ICAMS"))
list.of.vcfs <- ReadAndSplitStrelkaSBSVCFs(file)
SBS.vcf <- list.of.vcfs$SBS.vcfs[[1]]
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  annotated.SBS.vcf <- AnnotateSBSVCF(SBS.vcf, ref.genome = "hg19",
                                     trans.ranges = trans.ranges.GRCh37)
  PlotTransBiasGeneExpToPdf(annotated.SBS.vcf = annotated.SBS.vcf,
                           expression.data = gene.expression.data.HepG2,
                           Ensembl.gene.ID.col = "Ensembl.gene.ID",
                           expression.value.col = "TPM",
                           num.of.bins = 4,
```

```

    plot.type = c("C>A", "C>G", "C>T", "T>A", "T>C"),
    file = file.path(tempdir(), "test.pdf"))
}

```

---

ReadAndSplitMutectVCFs

*Read and split Mutect VCF files.*


---

## Description

Read and split Mutect VCF files.

## Usage

```
ReadAndSplitMutectVCFs(files, names.of.VCFs = NULL, tumor.col.names = NA)
```

## Arguments

<code>files</code>	Character vector of file paths to the Mutect VCF files.
<code>names.of.VCFs</code>	Optional. Character vector of names of the VCF files. The order of names in <code>names.of.VCFs</code> should match the order of VCF file paths in <code>files</code> . If <code>NULL</code> (default), this function will remove all of the path up to and including the last path separator (if any) in <code>files</code> and file paths without extensions (and the leading dot) will be used as the names of the VCF files.
<code>tumor.col.names</code>	Optional. Character vector of column names in VCFs which contain the tumor sample information. The order of names in <code>tumor.col.names</code> should match the order of VCFs specified in <code>files</code> . If <code>tumor.col.names</code> is equal to <code>NA</code> (default), this function will use the 10th column in all the VCFs to calculate VAFs. See <a href="#">GetMutectVAF</a> for more details.

## 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.
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) and rows for complex indels.
5. `multiple.alt` 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](#)

**Examples**

```
file <- c(system.file("extdata/Mutect-vcf",
                     "Mutect.GRCh37.vcf",
                     package = "ICAMS"))
list.of.vcfs <- ReadAndSplitMutectVCFs(file)
```

---

ReadAndSplitStrelkaSBSVCFs

*Read and split Strelka SBS VCF files.*


---

**Description**

Read and split Strelka SBS VCF files.

**Usage**

```
ReadAndSplitStrelkaSBSVCFs(files, names.of.VCFs = NULL)
```

**Arguments**

<code>files</code>	Character vector of file paths to the Strelka SBS VCF files.
<code>names.of.VCFs</code>	Optional. Character vector of names of the VCF files. The order of names in <code>names.of.VCFs</code> should match the order of VCF file paths in <code>files</code> . If <code>NULL</code> (default), this function will remove all of the path up to and including the last path separator (if any) in <code>files</code> and file paths without extensions (and the leading dot) will be used as the names of the 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` which contain rows in the input that did not represent SBSs or DBSs.
4. `multiple.alt` Rows with multiple alternate alleles (removed from `SBS.vcfs` etc.)

**See Also**

[StrelkaSBSVCFFilesToCatalog](#)

**Examples**

```
file <- c(system.file("extdata/Strelka-SBS-vcf",
                     "Strelka.SBS.GRCh37.vcf",
                     package = "ICAMS"))
list.of.vcfs <- ReadAndSplitStrelkaSBSVCFs(file)
```

---

ReadCatalog	<i>Read catalog.</i>
-------------	----------------------

---

## Description

Read a catalog in standardized format from path.

## Usage

```
ReadCatalog(  
  file,  
  ref.genome = NULL,  
  region = "unknown",  
  catalog.type = "counts",  
  strict = TRUE  
)
```

## Arguments

file	Path to a catalog on disk in the standardized format.
ref.genome	A ref.genome argument as described in <a href="#">ICAMS</a> .
region	region A character string designating a genomic region; see <a href="#">as.catalog</a> and <a href="#">ICAMS</a> .
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](#).

## Comments

To add or change attributes of the catalog, you can use function [attr](#).  
For example, `attr(catalog, "abundance") <- custom.abundance`.

## Note

In ID (small 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

```
file <- system.file("extdata",  
  "strelka.regress.cat.sbs.96.csv",  
  package = "ICAMS")  
catSBS96 <- ReadCatalog(file)
```

---

ReadStrelkaIDVCFs	<i>Read Strelka ID (small insertion and deletion) VCF files.</i>
-------------------	--

---

**Description**

Read Strelka ID (small insertion and deletion) VCF files.

**Usage**

```
ReadStrelkaIDVCFs(files, names.of.VCFs = NULL)
```

**Arguments**

<code>files</code>	Character vector of file paths to the Strelka ID VCF files.
<code>names.of.VCFs</code>	Optional. Character vector of names of the VCF files. The order of names in <code>names.of.VCFs</code> should match the order of VCF file paths in <code>files</code> . If <code>NULL</code> (default), this function will remove all of the path up to and including the last path separator (if any) in <code>files</code> and file paths without extensions (and the leading dot) will be used as the names of the VCF files.

**Value**

A list of data frames containing data lines of the VCF files.

**Note**

In ID (small 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**

```
file <- c(system.file("extdata/Strelka-ID-vcf",
                     "Strelka.ID.GRCh37.vcf",
                     package = "ICAMS"))
list.of.vcfs <- ReadStrelkaIDVCFs(file)
```

---

revc	<i>Reverse complement every string in string.vec.</i>
------	---

---

**Description**

Based on [reverseComplement](#). Handles IUPAC ambiguity codes but not "u" (uracil). (see <[https://en.wikipedia.org/wiki/Nucleic\\_acid\\_notation](https://en.wikipedia.org/wiki/Nucleic_acid_notation)>).

**Usage**

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



**Arguments**

`string.vec`      A character vector.

**Value**

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

**Examples**

```
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 (small insertion and deletion) catalog from Strelka ID VCF files*

---

**Description**

Create ID (small insertion and deletion) catalog from the Strelka ID VCFs specified by files

**Usage**

```
StrelkaIDVCFFilesToCatalog(
  files,
  ref.genome,
  region = "unknown",
  names.of.VCFs = NULL,
  flag.mismatches = 0
)
```

**Arguments**

<code>files</code>	Character vector of file paths to the Strelka ID VCF files.
<code>ref.genome</code>	A <code>ref.genome</code> argument as described in <a href="#">ICAMS</a> .
<code>region</code>	A character string designating a genomic region; see <a href="#">as.catalog</a> and <a href="#">ICAMS</a> .
<code>names.of.VCFs</code>	Optional. Character vector of names of the VCF files. The order of names in <code>names.of.VCFs</code> should match the order of VCF file paths in <code>files</code> . If <code>NULL</code> (default), this function will remove all of the path up to and including the last path separator (if any) in <code>files</code> and file paths without extensions (and the leading dot) will be used as the names of the VCF files.
<code>flag.mismatches</code>	Optional. If $> 0$ , then if there are mismatches to references in the ID (insertion/deletion) VCF, generate messages showing the mismatched rows and continue. Otherwise stop if there are mismatched rows. See <a href="#">AnnotateIDVCF</a> for more details.

**Details**

This function calls [VCFsToIDCatalogs](#)

**Value**

A list of two elements. 1st element is an S3 object containing an ID (small insertion and deletion) catalog with class "IndelCatalog". See [as.catalog](#) for more details. 2nd element is a list of further annotated VCFs.

**Note**

In ID (small 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**

```
file <- c(system.file("extdata/Strelka-ID-vcf",
                     "Strelka.ID.GRCh37.vcf",
                     package = "ICAMS"))
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  catID <- StrelkaIDVCFFilesToCatalog(file, ref.genome = "hg19",
                                     region = "genome")}
```

---

StrelkaIDVCFFilesToCatalogAndPlotToPdf

*Create ID (small insertion and deletion) catalog from Strelka ID VCF files and plot them to PDF*

---

**Description**

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

**Usage**

```
StrelkaIDVCFFilesToCatalogAndPlotToPdf(
  files,
  ref.genome,
  region = "unknown",
  names.of.VCFs = NULL,
  output.file = "",
  flag.mismatches = 0
)
```

**Arguments**

files	Character vector of file paths to the Strelka ID VCF files.
ref.genome	A ref.genome argument as described in <a href="#">ICAMS</a> .
region	A character string designating a genomic region; see <a href="#">as.catalog</a> and <a href="#">ICAMS</a> .

- `names.of.VCFs` Optional. Character vector of names of the VCF files. The order of names in `names.of.VCFs` should match the order of VCF file paths in `files`. If `NULL` (default), this function will remove all of the path up to and including the last path separator (if any) in `files` and file paths without extensions (and the leading dot) will be used as the names of the VCF files.
- `output.file` Optional. The base name of the PDF file to be produced; the file is ending in `catID.pdf`.
- `flag.mismatches` Optional. If  $> 0$ , then if there are mismatches to references in the ID (insertion/deletion) VCF, generate messages showing the mismatched rows and continue. Otherwise stop if there are mismatched rows. See [AnnotateIDVCF](#) for more details.

### Details

This function calls [StrelkaIDVCFFilesToCatalog](#) and [PlotCatalogToPdf](#)

### Value

A list whose first element is an ID (small insertion and deletion) catalog with its graph plotted to PDF with specified file name. The ID catalog has class attribute "IndelCatalog" added. See [as.catalog](#) for more details. The second element of the returned list is a list of further annotated VCFs.

### Note

In ID (small 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

```
file <- c(system.file("extdata/Strelka-ID-vcf",
                     "Strelka.ID.GRCh37.vcf",
                     package = "ICAMS"))
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  catID <-
    StrelkaIDVCFFilesToCatalogAndPlotToPdf(file, ref.genome = "hg19",
                                             region = "genome",
                                             output.file =
                                             file.path(tempdir(), "StrelkaID"))}
```

---

### StrelkaIDVCFFilesToZipFile

*Create a zip file which contains ID (small insertion and deletion) catalog and plot PDF from Strelka ID VCF files*

---

### Description

Create ID (small insertion and deletion) catalog from the Strelka ID VCFs specified by `dir`, save the catalog as CSV file, plot it to PDF and generate a zip archive of all the output files.

**Usage**

```
StrelkaIDVCFFilesToZipFile(
  dir,
  zipfile,
  ref.genome,
  region = "unknown",
  names.of.VCFs = NULL,
  base.filename = "",
  flag.mismatches = 0
)
```

**Arguments**

<code>dir</code>	Pathname of the directory which contains the Strelka ID VCF files. Each Strelka ID VCF <b>must</b> have a file extension ".vcf" (case insensitive) and share the <b>same</b> <code>ref.genome</code> and <code>region</code> .
<code>zipfile</code>	Pathname of the zip file to be created.
<code>ref.genome</code>	A <code>ref.genome</code> argument as described in <a href="#">ICAMS</a> .
<code>region</code>	A character string designating a genomic region; see <a href="#">as.catalog</a> and <a href="#">ICAMS</a> .
<code>names.of.VCFs</code>	Optional. Character vector of names of the VCF files. The order of names in <code>names.of.VCFs</code> should match the order of VCFs listed in <code>dir</code> . If <code>NULL</code> (default), this function will remove all of the path up to and including the last path separator (if any) in <code>dir</code> and file paths without extensions (and the leading dot) will be used as the names of the VCF files.
<code>base.filename</code>	Optional. The base name of the CSV and PDF file to be produced; the file is ending in <code>catID.csv</code> and <code>catID.pdf</code> respectively.
<code>flag.mismatches</code>	Optional. If $> 0$ , then if there are mismatches to references in the ID (insertion/deletion) VCF, generate messages showing the mismatched rows and continue. Otherwise stop if there are mismatched rows. See <a href="#">AnnotateIDVCF</a> for more details.

**Details**

This function calls [StrelkaIDVCFFilesToCatalog](#), [PlotCatalogToPdf](#), [WriteCatalog](#) and [zipr](#).

**Value**

A list of two elements. 1st element is an S3 object containing an ID (small insertion and deletion) catalog with class "IndelCatalog". See [as.catalog](#) for more details. 2nd element is a list of further annotated VCFs.

**Note**

In ID (small 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**

```
dir <- c(system.file("extdata/Strelka-ID-vcf",
```

```

        package = "ICAMS"))
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  catalogs <-
    StrelkaIDVCFFilesToZipFile(dir,
                              zipfile = file.path(tempdir(), "test.zip"),
                              ref.genome = "hg19",
                              region = "genome",
                              base.filename = "Strelka-ID")
  unlink(file.path(tempdir(), "test.zip"))}

```

---

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

## Usage

```

StrelkaSBSVCFFilesToCatalog(
  files,
  ref.genome,
  trans.ranges = NULL,
  region = "unknown",
  names.of.VCFs = NULL
)

```

## Arguments

<code>files</code>	Character vector of file paths to the Strelka SBS VCF files.
<code>ref.genome</code>	A <code>ref.genome</code> argument as described in <a href="#">ICAMS</a> .
<code>trans.ranges</code>	Optional. If <code>ref.genome</code> specifies one of the <a href="#">BSgenome</a> object <ol style="list-style-type: none"> <li><a href="#">BSgenome.Hsapiens.1000genomes.hs37d5</a></li> <li><a href="#">BSgenome.Hsapiens.UCSC.hg38</a></li> <li><a href="#">BSgenome.Mmusculus.UCSC.mm10</a></li> </ol> then the function will infer <code>trans.ranges</code> automatically. Otherwise, user will need to provide the necessary <code>trans.ranges</code> . Please refer to <a href="#">TranscriptRanges</a> for more details. If <code>is.null(trans.ranges)</code> do not add transcript range information.
<code>region</code>	A character string designating a genomic region; see <a href="#">as.catalog</a> and <a href="#">ICAMS</a> .
<code>names.of.VCFs</code>	Optional. Character vector of names of the VCF files. The order of names in <code>names.of.VCFs</code> should match the order of VCF file paths in <code>files</code> . If <code>NULL</code> (default), this function will remove all of the path up to and including the last path separator (if any) in <code>files</code> and file paths without extensions (and the leading dot) will be used as the names of the VCF files.

## 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` is not provided by user and cannot be inferred by ICAMS, SBS 192 and DBS 144 catalog will not be generated. Each catalog has attributes added. See [as.catalog](#) for more details.

**Comments**

To add or change attributes of the catalog, you can use function [attr](#).  
For example, `attr(catalog, "abundance") <- custom.abundance`.

**Note**

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

**Examples**

```
file <- c(system.file("extdata/Strelka-SBS-vcf",
                     "Strelka.SBS.GRCh37.vcf",
                     package = "ICAMS"))
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  catalogs <- StrelkaSBSVCFFilesToCatalog(file, ref.genome = "hg19",
                                          trans.ranges = trans.ranges.GRCh37,
                                          region = "genome")}
```

---

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

**Usage**

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

**Arguments**

<code>files</code>	Character vector of file paths to the Strelka SBS VCF files.
<code>ref.genome</code>	A <code>ref.genome</code> argument as described in <a href="#">ICAMS</a> .
<code>trans.ranges</code>	Optional. If <code>ref.genome</code> specifies one of the <a href="#">BSgenome</a> object



---

StrelkaSBSVCFFilesToZipFile

*Create a zip file which contains catalogs and plot PDFs from Strelka SBS VCF files*

---

## Description

Create 3 SBS catalogs (96, 192, 1536), 3 DBS catalogs (78, 136, 144) from the Strelka SBS VCFs specified by `dir`, save the catalogs as CSV files, plot them to PDF and generate a zip archive of all the output files.

## Usage

```
StrelkaSBSVCFFilesToZipFile(
  dir,
  zipfile,
  ref.genome,
  trans.ranges = NULL,
  region = "unknown",
  names.of.VCFs = NULL,
  base.filename = ""
)
```

## Arguments

<code>dir</code>	Pathname of the directory which contains the Strelka SBS VCF files. Each Strelka SBS VCF <b>must</b> have a file extension ".vcf" (case insensitive) and share the <b>same</b> <code>ref.genome</code> and <code>region</code> .
<code>zipfile</code>	Pathname of the zip file to be created.
<code>ref.genome</code>	A <code>ref.genome</code> argument as described in <a href="#">ICAMS</a> .
<code>trans.ranges</code>	Optional. If <code>ref.genome</code> specifies one of the <a href="#">BSgenome</a> object <ol style="list-style-type: none"> <li><a href="#">BSgenome.Hsapiens.1000genomes.hs37d5</a></li> <li><a href="#">BSgenome.Hsapiens.UCSC.hg38</a></li> <li><a href="#">BSgenome.Mmusculus.UCSC.mm10</a></li> </ol> then the function will infer <code>trans.ranges</code> automatically. Otherwise, user will need to provide the necessary <code>trans.ranges</code> . Please refer to <a href="#">TranscriptRanges</a> for more details. If <code>is.null(trans.ranges)</code> do not add transcript range information.
<code>region</code>	A character string designating a genomic region; see <a href="#">as.catalog</a> and <a href="#">ICAMS</a> .
<code>names.of.VCFs</code>	Optional. Character vector of names of the VCF files. The order of names in <code>names.of.VCFs</code> should match the order of VCFs listed in <code>dir</code> . If <code>NULL</code> (default), this function will remove all of the path up to and including the last path separator (if any) in <code>dir</code> and file paths without extensions (and the leading dot) will be used as the names of the VCF files.
<code>base.filename</code>	Optional. The base name of the CSV and PDF files to be produced; multiple files will be generated, each ending in <code>x.csv</code> or <code>x.pdf</code> , where <code>x</code> indicates the type of catalog.



**Details**

This function calls [StrelkaSBSVCFFilesToCatalog](#), [PlotCatalogToPdf](#), [WriteCatalog](#) and [zipr](#).

**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` is not provided by user and cannot be inferred by ICAMS, then SBS 192 and DBS 144 catalog will not be generated and plotted. Each catalog has attributes added. See [as.catalog](#) for more details.

**Comments**

To add or change attributes of the catalog, you can use function [attr](#).  
For example, `attr(catalog, "abundance") <- custom.abundance`.

**Note**

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

**Examples**

```
dir <- c(system.file("extdata/Strelka-SBS-vcf",
                    package = "ICAMS"))
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  catalogs <-
    StrelkaSBSVCFFilesToZipFile(dir,
                                zipfile = file.path(tempdir(), "test.zip"),
                                ref.genome = "hg19",
                                trans.ranges = trans.ranges.GRCh37,
                                region = "genome",
                                base.filename = "Strelka-SBS")
  unlink(file.path(tempdir(), "test.zip"))}
```

---

TranscriptRanges

*Transcript ranges data*


---

**Description**

Transcript ranges and strand information for a particular reference genome.

**Usage**

```
trans.ranges.GRCh37
```

```
trans.ranges.GRCh38
```

```
trans.ranges.GRCm38
```

## Format

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

An object of class `data.table` (inherits from `data.frame`) with 19083 rows and 6 columns.

An object of class `data.table` (inherits from `data.frame`) with 19096 rows and 6 columns.

An object of class `data.table` (inherits from `data.frame`) with 20325 rows and 6 columns.

## 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/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_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](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 Ensembl.gene.ID  gene.symbol
#    1      65419     71585      + ENSG00000186092      OR4F5
#    1     367640    368634      + ENSG00000235249      OR4F29
#    1     621059    622053      - ENSG00000284662      OR4F16
#    1     859308    879961      + ENSG00000187634      SAMD11
#    1     879583    894689      - ENSG00000188976      NOC2L
#    ...         ...         ...      ...         ...         ...
```

---

TransformCatalog	<i>Transform between counts and density spectrum catalogs and counts and density signature catalogs.</i>
------------------	--

---

## Description

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

## Usage

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

## Arguments

catalog	An SBS or DBS catalog as described in <a href="#">ICAMS</a> ; must <b>not</b> be an ID (small insertion and deletion) catalog.
target.ref.genome	A ref.genome argument as described in <a href="#">ICAMS</a> . If NULL, then defaults to the ref.genome attribute of catalog.
target.region	A region argument; see <a href="#">as.catalog</a> and <a href="#">ICAMS</a> . 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 <a href="#">as.catalog</a> . If NULL, then defaults to the catalog.type attribute of catalog.
target.abundance	A vector of counts, one for each source K-mer for mutations (e.g. for strand-agnostic single nucleotide substitutions in trinucleotide – i.e. 3-mer – context, one count each for ACA, ACC, ACG, ... TTT). See <a href="#">all.abundance</a> . If NULL, the function tries to infer target.abundance from the class of catalog and the value of the target.ref.genome, target.region, and target.catalog.type. If the target.abundance can be inferred and is different from a supplied non-NULL value of target.abundance, raise an error.

## Details

Only the following transformations are legal:

1. counts -> counts (deprecated, generates a warning; we strongly suggest that you work with densities if comparing spectra or signatures generated from data with different underlying abundances.)
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](#).

### Examples

```
file <- system.file("extdata",
                    "strelka.regress.cat.sbs.96.csv",
                    package = "ICAMS")
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  catSBS96.counts <- ReadCatalog(file, ref.genome = "hg19",
                                region = "genome",
                                catalog.type = "counts")
  catSBS96.density <- TransformCatalog(catSBS96.counts,
                                       target.ref.genome = "hg19",
                                       target.region = "genome",
                                       target.catalog.type = "density")}
```

---

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 than DBSs.

### Usage

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

**Arguments**

<code>list.of.DBS.vcfs</code>	List of in-memory data frames of pure DBS mutations – no SBS or 3+BS mutations. The list names will be the sample ids in the output catalog.
<code>ref.genome</code>	A <code>ref.genome</code> argument as described in <a href="#">ICAMS</a> .
<code>trans.ranges</code>	Optional. If <code>ref.genome</code> specifies one of the <a href="#">BSgenome</a> object <ol style="list-style-type: none"> <li><a href="#">BSgenome.Hsapiens.1000genomes.hs37d5</a></li> <li><a href="#">BSgenome.Hsapiens.UCSC.hg38</a></li> <li><a href="#">BSgenome.Mmusculus.UCSC.mm10</a></li> </ol> then the function will infer <code>trans.ranges</code> automatically. Otherwise, user will need to provide the necessary <code>trans.ranges</code> . Please refer to <a href="#">TranscriptRanges</a> for more details. If <code>is.null(trans.ranges)</code> do not add transcript range information.
<code>region</code>	A character string designating a genomic region; see <a href="#">as.catalog</a> and <a href="#">ICAMS</a> .

**Value**

A list of 3 DBS catalogs, one each for 78, 144, 136: `catDBS78` `catDBS144` `catDBS136`. If `trans.ranges` is not provided by user and cannot be inferred by [ICAMS](#), DBS 144 catalog will not be generated. Each catalog has attributes added. See [as.catalog](#) for more details.

**Comments**

To add or change attributes of the catalog, you can use function [attr](#).  
For example, `attr(catalog, "abundance") <- custom.abundance`.

**Note**

DBS 144 catalog only contains mutations in transcribed regions.

**Examples**

```
file <- c(system.file("extdata/Mutect-vcf",
                     "Mutect.GRCh37.vcf",
                     package = "ICAMS"))
list.of.DBS.vcfs <- ReadAndSplitMutectVCFs(file)$DBS
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  catalogs.DBS <- VCFsToIDCatalogs(list.of.DBS.vcfs, ref.genome = "hg19",
                                   trans.ranges = trans.ranges.GRCh37,
                                   region = "genome")}
```

---

VCFsToIDCatalogs

---

*Create ID (small insertion and deletion) catalog from ID VCFs*


---

**Description**

Create ID (small insertion and deletion) catalog from ID VCFs

**Usage**

```
VCFsToIDCatalogs(
  list.of.vcfs,
  ref.genome,
  region = "unknown",
  flag.mismatches = 0
)
```

**Arguments**

<code>list.of.vcfs</code>	List of in-memory VCFs. The list names will be the sample ids in the output catalog.
<code>ref.genome</code>	A <code>ref.genome</code> argument as described in <a href="#">ICAMS</a> .
<code>region</code>	A character string acting as a region identifier, one of "genome", "exome".
<code>flag.mismatches</code>	Optional. If > 0, then if there are mismatches to references in the ID (insertion/deletion) VCF, generate messages showing the mismatched rows and continue. Otherwise stop if there are mismatched rows. See <a href="#">AnnotateIDVCF</a> for more details.

**Value**

A list of two elements. 1st element is an S3 object containing an ID (small insertion and deletion) catalog with class "IndelCatalog". See [as.catalog](#) for more details. 2nd element is a list of further annotated VCFs.

**Note**

In ID (small 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**

```
file <- c(system.file("extdata/Strelka-ID-vcf/",
                     "Strelka.ID.GRCh37.vcf",
                     package = "ICAMS"))
list.of.ID.vcfs <- ReadStrelkaIDVCFs(file)
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5",
  quietly = TRUE)) {
  catID <- VCFsToIDCatalogs(list.of.ID.vcfs, ref.genome = "hg19",
    region = "genome")}
```

---

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.

**Usage**

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

**Arguments**

<code>list.of.SBS.vcfs</code>	List of in-memory data frames of pure SBS mutations – no DBS or 3+BS mutations. The list names will be the sample ids in the output catalog.
<code>ref.genome</code>	A <code>ref.genome</code> argument as described in <a href="#">ICAMS</a> .
<code>trans.ranges</code>	Optional. If <code>ref.genome</code> specifies one of the <a href="#">BSgenome</a> object <ol style="list-style-type: none"> <li>1. <a href="#">BSgenome.Hsapiens.1000genomes.hs37d5</a></li> <li>2. <a href="#">BSgenome.Hsapiens.UCSC.hg38</a></li> <li>3. <a href="#">BSgenome.Mmusculus.UCSC.mm10</a></li> </ol> then the function will infer <code>trans.ranges</code> automatically. Otherwise, user will need to provide the necessary <code>trans.ranges</code> . Please refer to <a href="#">TranscriptRanges</a> for more details. If <code>is.null(trans.ranges)</code> do not add transcript range information.
<code>region</code>	A character string designating a genomic region; see <a href="#">as.catalog</a> and <a href="#">ICAMS</a> .

**Value**

A list of 3 SBS catalogs, one each for 96, 192, 1536: `catSBS96` `catSBS192` `catSBS1536`. If `trans.ranges` is not provided by user and cannot be inferred by [ICAMS](#), SBS 192 catalog will not be generated. Each catalog has attributes added. See [as.catalog](#) for more details.

**Comments**

To add or change attributes of the catalog, you can use function [attr](#). For example, `attr(catalog, "abundance") <- custom.abundance`.

**Note**

SBS 192 catalogs only contain mutations in transcribed regions.

**Examples**

```
file <- c(system.file("extdata/Mutect-vcf",
  "Mutect.GRCh37.vcf",
  package = "ICAMS"))
list.of.SBS.vcfs <- ReadAndSplitMutectVCFs(file)$SBS
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  catalogs.SBS <- VCFsToSBSCatalogs(list.of.SBS.vcfs, ref.genome = "hg19",
    trans.ranges = trans.ranges.GRCh37,
    region = "genome")}
```

---

WriteCatalog	<i>Write a catalog</i>
--------------	------------------------

---

**Description**

Write a catalog to a file.

**Usage**

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

**Arguments**

catalog	A catalog as defined in <a href="#">ICAMS</a> ; see also <a href="#">as.catalog</a> .
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](#).

**Note**

In ID (small 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**

```
file <- system.file("extdata",  
                    "strelka.regress.cat.sbs.96.csv",  
                    package = "ICAMS")  
catSBS96 <- ReadCatalog(file)  
WriteCatalog(catSBS96, file = file.path(tempdir(), "catSBS96.csv"))
```



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