

# Package ‘ICAMS’

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**Type** Package

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

**Version** 2.3.10

**Author** Steve Rozen, Nanhai Jiang, Arnoud Boot, Mo Liu, Yang Wu

**Maintainer** Steve Rozen <steverozen@gmail.com>

**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> and "Characterization of colibactin-associated mutational signature in an Asian oral squamous cell carcinoma and in other mucosal tumor types", Genome Research 2020 <doi:10.1101/gr.255620.119>. '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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**URL** <https://github.com/steverozen/ICAMS>

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

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**biocViews**

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testthat

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<i>all.abundance</i>	<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, name.of.VCF = NULL)
```

## Arguments

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

## 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.s1.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.DB.S.vcf <- 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,
  name.of.VCF = NULL,
  suppress.discarded.variants.warnings = TRUE
)
```

### Arguments

ID.vcf	An in-memory ID (insertion/deletion) VCF as a data.frame. This function expects that there is a "context base" to the left, for example REF = ACG, ALT = A (deletion of CG) or REF = A, ALT = ACC (insertion of CC).
ref.genome	A ref.genome argument as described in <a href="#">ICAMS</a> .
flag.mismatches	Deprecated. If there are ID variants whose REF do not match the extracted sequence from ref.genome, the function will automatically discard these variants. See element discarded.variants in the return value for more details.
name.of.VCF	Name of the VCF file.
suppress.discarded.variants.warnings	Logical. Whether to suppress warning messages showing information about the discarded variants. Default is TRUE.

### Value

A list of elements:

- annotated.vcf: The original VCF data frame with two new columns added to the input data frame:
  - seq.context: The sequence embedding the variant.
  - seq.context.width: The width of seq.context to the left.
- discarded.variants: **Non-NULL only if** there are variants that were excluded from the analysis. See the added extra column discarded.reason for more details.

**Examples**

```
file <- c(system.file("extdata/Strelka-ID-vcf/",
                    "Strelka.ID.GRCh37.s1.vcf",
                    package = "ICAMS"))
ID.vcf <- ReadStrelkaIDVCFs(file)[[1]]
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  list <- AnnotateIDVCF(ID.vcf, ref.genome = "hg19")
  annotated.ID.vcf <- list$annotated.vcf}
```

---

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, name.of.VCF = NULL)
```

**Arguments**

SBS.vcf	An in-memory SBS VCF as a data.frame.
ref.genome	A ref.genome argument as described in <a href="#">ICAMS</a> .
trans.ranges	Optional. If ref.genome specifies one of the <a href="#">BSgenome</a> object <ol style="list-style-type: none"> <li>1. BSgenome.Hsapiens.1000genomes.hs37d5</li> <li>2. BSgenome.Hsapiens.UCSC.hg38</li> <li>3. BSgenome.Mmusculus.UCSC.mm10</li> </ol>

then the function will infer trans.ranges automatically. Otherwise, user will need to provide the necessary trans.ranges. Please refer to [TranscriptRanges](#) for more details. If is.null(trans.ranges) do not add transcript range information.

name.of.VCF	Name of the VCF file.
-------------	-----------------------

**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.s1.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 infer.rownames is FALSE then 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> . If the catalog type is a stranded catalog type (SBS192 or DBS144), region = "genome" will be silently converted to "transcript".
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, <b>be sure the order of rows is correct</b> .

**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**

context	The deleted sequence plus ample surrounding sequence on each side (at least as long as del.seq).
del.seq	The deleted sequence in context.
pos	The position of del.sequence in context.
trace	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 [FindDelMH](#)).

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 [FindDelMH](#) 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
```

**Format**

A list of character vectors indicating the standard orders of row names in catalogs.  
An object of class `list` of length 8.

**ID classification**

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 ID (small insertion and deletion) mutation classification.

See the documentation for `Canonicalize1Del` which 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 `FindDelMH`).

See the code for unexported function `CanonicalizeID` and the functions it calls for handling of insertions.

**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.
```

---

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
....
```

=&gt;

```
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.

**ID classification**

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 ID (small insertion and deletion) mutation classification.

See the documentation for [Canonicalize1Del](#) which 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 [FindDelMH](#)).

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

**Examples**

```
# GAGAGG[CTAGAA]CTAGTT
#      ----  ----
FindDelMH("GGAGAGGCTAGAACTAGTTAAAA", "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 <code>rep.unit.seq</code> at position <code>pos</code>
<code>rep.unit.seq</code>	A substring of <code>context</code> at <code>pos</code> to <code>pos + nchar(rep.unit.seq) - 1</code> , which is the repeat unit sequence.
<code>pos</code>	The position of <code>rep.unit.seq</code> in <code>context</code> .

**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.

**ID classification**

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 ID (small insertion and deletion) mutation classification.

See the documentation for [Canonicalize1Del](#) which 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 [FindDelMH](#)).

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

**Examples**

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

---

GeneExpressionData      *Example gene expression data from two cell lines*

---

**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
# ENSG00000000003    TSPAN6    6007    33.922648455
# ENSG00000000005      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)
```

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

### Arguments

<code>vcf</code>	Said VCF as a <code>data.frame</code> .
<code>name.of.VCF</code>	Name of the VCF file.
<code>tumor.col.name</code>	Optional. Only applicable to <b>Mutect</b> VCF. Name of the column in <b>Mutect</b> VCF which contains the tumor sample information. It <b>must</b> have quotation marks. If <code>tumor.col.name</code> is equal to <code>NA</code> (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.s1.vcf",  
                    package = "ICAMS"))  
MakeDataFrameFromVCF <- getFromNamespace("MakeDataFrameFromVCF", "ICAMS")  
df <- MakeDataFrameFromVCF(file)  
df1 <- GetStrelkaVAF(df)
```



## 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> and "Characterization of colibactin-associated mutational signature in an Asian oral squamous cell carcinoma and in other mucosal tumor types", *Genome Research* 2020, <https://doi.org/10.1101/gr.255620.119>. "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. [VCFsToCatalogs](#) creates 3 SBS catalogs (96, 192, 1536), 3 DBS catalogs (78, 136, 144) and ID (small insertion and deletion) catalog from the VCFs. It has more general usage with functionalities overlapping with the three functions below. For example, it is the same as [MutectVCFFilesToCatalog](#) when `variant.caller = "mutect"`.
2. [StrelkaSBSVCFFilesToCatalog](#) creates 3 SBS catalogs (96, 192, 1536) and 3 DBS catalogs (78, 136, 144) from the Strelka SBS VCFs.
3. [StrelkaIDVCFFilesToCatalog](#) creates an ID (small insertion and deletion) catalog from the Strelka ID VCFs.
4. [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. [VCFsToCatalogsAndPlotToPdf](#) creates all types of SBS, DBS and ID catalogs from VCFs and plots the catalogs. It has more general usage with functionalities overlapping with the three functions below. For example, it is the same as [MutectVCFFilesToCatalogAndPlotToPdf](#) when `variant.caller = "mutect"`.

2. [StrelkaSBSVCFFilesToCatalogAndPlotToPdf](#) creates all type of SBS and DBS catalogs from Strelka SBS VCFs and plots the catalogs.
3. [StrelkaIDVCFFilesToCatalogAndPlotToPdf](#) creates an ID (small insertion and deletion) catalog from Strelka ID VCFs and plot it.
4. [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. [VCFsToZipFile](#) creates a zip file which contains SBS, DBS and ID catalogs and plot PDFs from VCF files. It has more general usage with functionalities overlapping with the three functions below. For example, it is the same as [MutectVCFFilesToZipFile](#) when `variant.caller = "mutect"`.
2. [StrelkaSBSVCFFilesToZipFile](#) creates a zip file which contains SBS and DBS catalogs and plot PDFs from Strelka SBS VCF files.
3. [StrelkaIDVCFFilesToZipFile](#) creates a zip file which contains ID (small insertion and deletion) catalog and plot PDF from Strelka ID VCF files.
4. [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/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. 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,
  return.annotated.vcfs = FALSE,
  suppress.discarded.variants.warnings = TRUE
)
```

## 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><code>BSgenome.Hsapiens.1000genomes.hs37d5</code></li> <li><code>BSgenome.Hsapiens.UCSC.hg38</code></li> <li><code>BSgenome.Mmusculus.UCSC.mm10</code></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 [GetMutectVAF](#) for more details.

`flag.mismatches`

Deprecated. If there are ID variants whose REF do not match the extracted sequence from `ref.genome`, the function will automatically discard these variants and an element `discarded.variants` will appear in the return value. See [AnnotateIDVCF](#) for more details.

`return.annotated.vcfs`

Logical. Whether to return the annotated VCFs with additional columns showing mutation class for each variant. Default is FALSE.

`suppress.discarded.variants.warnings`

Logical. Whether to suppress warning messages showing information about the discarded variants. Default is TRUE.

## Details

This function calls [VCFsToSBSCatalogs](#), [VCFsToDBSCatalogs](#) and [VCFsToIDCatalogs](#)

## Value

A list containing the following objects:

- `catSBS96`, `catSBS192`, `catSBS1536`: Matrix of 3 SBS catalogs (one each for 96, 192, and 1536).
- `catDBS78`, `catDBS136`, `catDBS144`: Matrix of 3 DBS catalogs (one each for 78, 136, and 144).
- `catID`: Matrix of ID (small insertion and deletion) catalog.
- `discarded.variants`: **Non-NULL only if** there are variants that were excluded from the analysis. See the added extra column `discarded.reason` for more details.
- `annotated.vcfs`: **Non-NULL only if** `return.annotated.vcfs = TRUE`. A list of elements:
  - SBS: SBS VCF annotated by [AnnotateSBSVCF](#) with three new columns `SBS96.class`, `SBS192.class` and `SBS1536.class` showing the mutation class for each SBS variant.
  - DBS: DBS VCF annotated by [AnnotateDBSVCF](#) with three new columns `DBS78.class`, `DBS136.class` and `DBS144.class` showing the mutation class for each DBS variant.
  - ID: ID VCF annotated by [AnnotateIDVCF](#) with one new column `ID.class` showing the mutation class for each ID variant.

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.

## ID classification

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 ID (small insertion and deletion) mutation classification.

See the documentation for [Canonicalize1Del](#) which 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 [FindDelMH](#)).

See the code for unexported function `CanonicalizeID` and the functions it calls for handling of insertions.

### 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+.

### Comments

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

### Examples

```
file <- c(system.file("extdata/Mutect-vcf",
                    "Mutect.GRCh37.s1.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,
  return.annotated.vcfs = FALSE,
  suppress.discarded.variants.warnings = TRUE
)
```

**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 <ul style="list-style-type: none"> <li>1. <code>BSgenome.Hsapiens.1000genomes.hs37d5</code></li> <li>2. <code>BSgenome.Hsapiens.UCSC.hg38</code></li> <li>3. <code>BSgenome.Mmusculus.UCSC.mm10</code></li> </ul> 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>	Deprecated. If there are ID variants whose REF do not match the extracted sequence from <code>ref.genome</code> , the function will automatically discard these variants and an element <code>discarded.variants</code> will appear in the return value. See <a href="#">AnnotateIDVCF</a> for more details.
<code>return.annotated.vcfs</code>	Logical. Whether to return the annotated VCFs with additional columns showing mutation class for each variant. Default is <code>FALSE</code> .
<code>suppress.discarded.variants.warnings</code>	Logical. Whether to suppress warning messages showing information about the discarded variants. Default is <code>TRUE</code> .

**Details**

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

**Value**

A list containing the following objects:



- `catSBS96`, `catSBS192`, `catSBS1536`: Matrix of 3 SBS catalogs (one each for 96, 192, and 1536).
- `catDBS78`, `catDBS136`, `catDBS144`: Matrix of 3 DBS catalogs (one each for 78, 136, and 144).
- `catID`: Matrix of ID (small insertion and deletion) catalog.
- `discarded.variants`: **Non-NULL only if** there are variants that were excluded from the analysis. See the added extra column `discarded.reason` for more details.
- `annotated.vcfs`: **Non-NULL only if** `return.annotated.vcfs = TRUE`. A list of elements:
  - SBS: SBS VCF annotated by `AnnotateSBSVCF` with three new columns `SBS96.class`, `SBS192.class` and `SBS1536.class` showing the mutation class for each SBS variant.
  - DBS: DBS VCF annotated by `AnnotateDBSVCF` with three new columns `DBS78.class`, `DBS136.class` and `DBS144.class` showing the mutation class for each DBS variant.
  - ID: ID VCF annotated by `AnnotateIDVCF` with one new column `ID.class` showing the mutation class for each ID variant.

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.

## ID classification

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 ID (small insertion and deletion) mutation classification.

See the documentation for `Canonicalize1Del` which 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 `FindDelMH`).

See the code for unexported function `CanonicalizeID` and the functions it calls for handling of insertions.

## 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+.

## Comments

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

## Examples

```
file <- c(system.file("extdata/Mutect-vcf",
                     "Mutect.GRCh37.s1.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,
  return.annotated.vcfs = FALSE,
  suppress.discarded.variants.warnings = TRUE
)

```

### Arguments

<code>dir</code>	Pathname of the directory which contains <b>only</b> 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 region.
<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. <code>BSgenome.Hsapiens.1000genomes.hs37d5</code></li> <li>2. <code>BSgenome.Hsapiens.UCSC.hg38</code></li> <li>3. <code>BSgenome.Mmusculus.UCSC.mm10</code></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>	Deprecated. If there are ID variants whose REF do not match the extracted sequence from <code>ref.genome</code> , the function will automatically discard these variants and an element <code>discarded.variants</code> will appear in the return value. See <a href="#">AnnotateIDVCF</a> for more details.
<code>return.annotated.vcfs</code>	Logical. Whether to return the annotated VCFs with additional columns showing mutation class for each variant. Default is <code>FALSE</code> .
<code>suppress.discarded.variants.warnings</code>	Logical. Whether to suppress warning messages showing information about the discarded variants. Default is <code>TRUE</code> .

## Details

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

## Value

A list containing the following objects:

- `catSBS96`, `catSBS192`, `catSBS1536`: Matrix of 3 SBS catalogs (one each for 96, 192, and 1536).
- `catDBS78`, `catDBS136`, `catDBS144`: Matrix of 3 DBS catalogs (one each for 78, 136, and 144).
- `catID`: Matrix of ID (small insertion and deletion) catalog.
- `discarded.variants`: **Non-NULL only if** there are variants that were excluded from the analysis. See the added extra column `discarded.reason` for more details.
- `annotated.vcfs`: **Non-NULL only if** `return.annotated.vcfs = TRUE`. A list of elements:
  - SBS: SBS VCF annotated by [AnnotateSBSVCF](#) with three new columns `SBS96.class`, `SBS192.class` and `SBS1536.class` showing the mutation class for each SBS variant.
  - DBS: DBS VCF annotated by [AnnotateDBSVCF](#) with three new columns `DBS78.class`, `DBS136.class` and `DBS144.class` showing the mutation class for each DBS variant.

- ID: ID VCF annotated by `AnnotateIDVCF` with one new column `ID.class` showing the mutation class for each ID variant.

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.

## ID classification

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 ID (small insertion and deletion) mutation classification.

See the documentation for `Canonicalize1Del` which 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 `FindDelMH`).

See the code for unexported function `CanonicalizeID` and the functions it calls for handling of insertions.

## 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+.

## Comments

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

## 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	<i>Plot <b>one</b> spectrum or signature</i>
-------------	--

---

### Description

Plot the spectrum of **one** sample or plot **one** signature. The type of graph is based on 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,
  xlabel = NULL,
  ylabel = 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. catalog can also be a numeric matrix, numeric data.frame, or a vector denoting the mutation <b>counts</b> , but <b>must</b> be in the correct row order used in <a href="#">ICAMS</a> . See <a href="#">CatalogRowOrder</a> for more details. If catalog is a vector, it will be converted to a 1-column matrix with rownames taken from the element names of the vector and with column name "Unknown".
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	Has the usual meaning. Taken from par("cex") by default. Only implemented for SBS96Catalog, SBS192Catalog and DBS144Catalog.
grid	A logical value indicating whether to draw grid lines. Only implemented for SBS96Catalog, DBS78Catalog, IndelCatalog.
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, DBS78Catalog, IndelCatalog.
xlabel	A logical value indicating whether to draw x axis labels. Only implemented for SBS96Catalog, DBS78Catalog, IndelCatalog. If FALSE then plot x axis tick marks for SBS96Catalog; set par(tck = 0) to suppress.

ylabels	A logical value indicating whether to draw y axis labels. Only implemented for SBS96Catalog, DBS78Catalog, IndelCatalog.
yylim	Has the usual meaning. Only implemented for SBS96Catalog and IndelCatalog.

### Value

An **invisible** list whose first element is a logic value indicating whether the plot is successful. For SBS96Catalog, SBS192Catalog, DBS78Catalog, DBS144Catalog and IndelCatalog, 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 and plot.SBS12 = TRUE, the list will have an additional element which is a list containing the strand bias statistics.

### Comments

For **SBS192Catalog** with "counts" catalog.type and non-NULL abundance and plot.SBS12 = TRUE, 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 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,
  ylabels = 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. catalog can also be a numeric matrix, numeric data.frame, or a vector denoting the mutation <b>counts</b> , but <b>must</b> be in the correct row order used in <a href="#">ICAMS</a> . See <a href="#">CatalogRowOrder</a> for more details. If catalog is a vector, it will be converted to a 1-column matrix with rownames taken from the element names of the vector and with column name "Unknown".
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	Has the usual meaning. A default value has been used by the program internally. Only implemented for SBS96Catalog, SBS192Catalog and DBS144Catalog.
grid	A logical value indicating whether to draw grid lines. Only implemented for SBS96Catalog, DBS78Catalog, IndelCatalog.
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, DBS78Catalog, IndelCatalog.
xlabels	A logical value indicating whether to draw x axis labels. Only implemented for SBS96Catalog, DBS78Catalog, IndelCatalog. If FALSE then plot x axis tick marks for SBS96Catalog; set par(tck = 0) to suppress.
ylabels	A logical value indicating whether to draw y axis labels. Only implemented for SBS96Catalog, DBS78Catalog, IndelCatalog.
ylim	Has the usual meaning. Only implemented for SBS96Catalog and IndelCatalog.

**Value**

An **invisible** list whose first element is a logic value indicating whether the plot is successful. For **SBS192Catalog** with "counts" catalog.type and non-null abundance and plot.SBS12 = TRUE, 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 and `plot.SBS12 = TRUE`, 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 [AnnotateSBSVCF](#). It **must** 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.
ymax	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.s1.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

annotated.SBS.vcf	An SBS VCF annotated by <a href="#">AnnotateSBSVCF</a> . It <b>must</b> have transcript range information added.
file	The name of output file.
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 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.
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.

**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.s1.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,
  suppress.discarded.variants.warnings = TRUE
)
```

**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.
<code>suppress.discarded.variants.warnings</code>	Logical. Whether to suppress warning messages showing information about the discarded variants. Default is <code>TRUE</code> .

**Value**

A list containing the following objects:

- `SBS`: List of VCFs with only single base substitutions.
- `DBS`: List of VCFs with only doublet base substitutions as called by Mutect.
- `ID`: List of VCFs with only small insertions and deletions.
- `discarded.variants`: **Non-NULL only if** there are variants that were excluded from the analysis. See the added extra column `discarded.reason` for more details.

**See Also**

[MutectVCFFilesToCatalog](#)

**Examples**

```
file <- c(system.file("extdata/Mutect-vcf",
                    "Mutect.GRCh37.s1.vcf",
                    package = "ICAMS"))
list.of.vcfs <- ReadAndSplitMutectVCFs(file)
```

---

ReadAndSplitStrelkaSBSVCFs

*Read and split Strelka SBS VCF files*

---

**Description**

The function will find and merge adjacent SBS pairs into DBS if their VAFs are very similar. The default threshold value for VAF is 0.02.

**Usage**

```
ReadAndSplitStrelkaSBSVCFs(
  files,
  names.of.VCFs = NULL,
  suppress.discarded.variants.warnings = TRUE
)
```

**Arguments**

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

`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.

`suppress.discarded.variants.warnings` Logical. Whether to suppress warning messages showing information about the discarded variants. Default is `TRUE`.

**Value**

A list of elements as follows:

- `SBS.vcfs`: List of data.frames of pure SBS mutations – no DBS or 3+BS mutations.
- `DBS.vcfs`: List of data.frames of pure DBS mutations – no SBS or 3+BS mutations.
- `discarded.variants`: **Non-NULL only** if there are variants that were excluded from the analysis. See the added extra column `discarded.reason` for more details.

**See Also**

[StrelkaSBSVCFFilesToCatalog](#)

**Examples**

```
file <- c(system.file("extdata/Strelka-SBS-vcf",
                     "Strelka.SBS.GRCh37.s1.vcf",
                     package = "ICAMS"))
list.of.vcfs <- ReadAndSplitStrelkaSBSVCFs(file)
```

---

ReadAndSplitVCFs

*Read and split VCF files*

---

**Description**

Read and split VCF files

**Usage**

```

ReadAndSplitVCFs(
  files,
  variant.caller = "unknown",
  num.of.cores = 1,
  names.of.VCFs = NULL,
  tumor.col.names = NA,
  filter.status = NULL,
  get.vaf.function = NULL,
  ...,
  max.vaf.diff = 0.02,
  suppress.discarded.variants.warnings = TRUE
)

```

**Arguments**

<code>files</code>	Character vector of file paths to the VCF files.
<code>variant.caller</code>	Name of the variant caller that produces the VCF, can be either "strelka", "mutect", "freebayes" or "unknown". This information is needed to calculate the VAFs (variant allele frequencies). If variant caller is "unknown"(default) and <code>get.vaf.function</code> is NULL, then VAF and read depth will be NAs. If variant caller is "mutect", do <b>not</b> merge SBSs into DBS.
<code>num.of.cores</code>	The number of cores to use. Not available on Windows unless <code>num.of.cores = 1</code> .
<code>names.of.VCFs</code>	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 NULL(default), this function will remove all of the path up to and including the last path separator (if any) 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. Only applicable to <b>Mutect</b> VCFs. Character vector of column names in <b>Mutect</b> VCFs which contain the tumor sample information. The order of names in <code>tumor.col.names</code> should match the order of <b>Mutect</b> VCFs specified in <code>files</code> . If <code>tumor.col.names</code> is equal to NA(default), this function will use the 10th column in all the <b>Mutect</b> VCFs to calculate VAFs. See <a href="#">GetMutectVAF</a> for more details.
<code>filter.status</code>	The status indicating a variant has passed all filters. An example would be "PASS". Variants which don't have the specified <code>filter.status</code> in the FILTER column in VCF will be removed. If NULL(default), no variants will be removed from the original VCF.
<code>get.vaf.function</code>	Optional. Only applicable when <code>variant.caller</code> is " <b>unknown</b> ". Function to calculate VAF(variant allele frequency) and read depth information from original VCF. See <a href="#">GetMutectVAF</a> as an example. If NULL(default) and <code>variant.caller</code> is "unknown", then VAF and read depth will be NAs.
<code>...</code>	Optional arguments to <code>get.vaf.function</code> .

- `max.vaf.diff` **Not** applicable if `variant.caller = "mutect"`. The maximum difference of VAF, default value is 0.02. If the absolute difference of VAFs for adjacent SBSs is bigger than `max.vaf.diff`, then these adjacent SBSs are likely to be "merely" asynchronous single base mutations, opposed to a simultaneous doublet mutation or variants involving more than two consecutive bases.
- `suppress.discarded.variants.warnings`  
Logical. Whether to suppress warning messages showing information about the discarded variants. Default is TRUE.

**Value**

A list containing the following objects:

- `SBS`: List of VCFs with only single base substitutions.
- `DBS`: List of VCFs with only doublet base substitutions.
- `ID`: List of VCFs with only small insertions and deletions.
- `discarded.variants`: **Non-NULL only if** there are variants that were excluded from the analysis. See the added extra column `discarded.reason` for more details.

**See Also**

[VCFsToCatalogs](#)

**Examples**

```
file <- c(system.file("extdata/Mutect-vcf",
                    "Mutect.GRCh37.s1.vcf",
                    package = "ICAMS"))
list.of.vcfs <- ReadAndSplitVCFs(file, variant.caller = "mutect")
```

---

ReadCatalog

*Read catalog*

---

**Description**

Read a catalog in standardized format from path.

**Usage**

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

**Arguments**

file	Path to a catalog on disk in a standardized format. The recognized formats are: <ul style="list-style-type: none"> <li>• ICAMS formatted SBS96, SBS192, SBS1536, DBS78, DBS136, DBS144, ID (see <a href="#">CatalogRowOrder</a>).</li> <li>• SigProfiler-formatted SBS96, DBS78 and ID83 catalogs; see <a href="https://github.com/AlexandrovLab/SigProfilerExtractor">https://github.com/AlexandrovLab/SigProfilerExtractor</a>.</li> <li>• COSMIC-formatted SBS96, SBS192 (a.k.a. TSB192), DBS78, ID83 catalogs; see <a href="https://cancer.sanger.ac.uk/signatures/">https://cancer.sanger.ac.uk/signatures/</a>.</li> </ul>
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	Ignored and deprecated.
stop.on.error	If TRUE, call stop on error; otherwise return a 1-column matrix of NA's with the attribute "error" containing error information. The number of rows may not be the correct number for the expected catalog type.

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

---

## 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 VCF files.
<code>names.of.VCFs</code>	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.s1.vcf",  
                    package = "ICAMS"))  
list.of.vcfs <- ReadStrelkaIDVCFs(file)
```

revc *Reverse complement every string in string.vec*

---

### 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,
  return.annotated.vcfs = FALSE,
  suppress.discarded.variants.warnings = TRUE
)
```

**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>	Deprecated. If there are ID variants whose REF do not match the extracted sequence from <code>ref.genome</code> , the function will automatically discard these variants and an element <code>discarded.variants</code> will appear in the return value. See <a href="#">AnnotateIDVCF</a> for more details.
<code>return.annotated.vcfs</code>	Logical. Whether to return the annotated VCFs with additional columns showing mutation class for each variant. Default is <code>FALSE</code> .
<code>suppress.discarded.variants.warnings</code>	Logical. Whether to suppress warning messages showing information about the discarded variants. Default is <code>TRUE</code> .

**Details**

This function calls [VCFsToIDCatalogs](#)

**Value**

A **list** of elements:

- `catalog`: The ID (small insertion and deletion) catalog with attributes added. See [as.catalog](#) for more details.
- `discarded.variants`: **Non-NULL only if** there are variants that were excluded from the analysis. See the added extra column `discarded.reason` for more details.

- annotated.vcfs: **Non-NULL only if** return.annotated.vcfs = TRUE. A list of data frames which contain the original VCF's ID mutation rows with three additional columns seq.context.width, seq.context and ID.class added. The category assignment of each ID mutation in VCF can be obtained from ID.class column.

### ID classification

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 ID (small insertion and deletion) mutation classification.

See the documentation for [Canonicalize1Del](#) which 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 [FindDelMH](#)).

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

### 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.s1.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,
return.annotated.vcfs = FALSE,
suppress.discarded.variants.warnings = TRUE
)

```

## 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>output.file</code>	Optional. The base name of the PDF file to be produced; the file is ending in <code>catID.pdf</code> .
<code>flag.mismatches</code>	Deprecated. If there are ID variants whose REF do not match the extracted sequence from <code>ref.genome</code> , the function will automatically discard these variants and an element <code>discarded.variants</code> will appear in the return value. See <a href="#">AnnotateIDVCF</a> for more details.
<code>return.annotated.vcfs</code>	Logical. Whether to return the annotated VCFs with additional columns showing mutation class for each variant. Default is <code>FALSE</code> .
<code>suppress.discarded.variants.warnings</code>	Logical. Whether to suppress warning messages showing information about the discarded variants. Default is <code>TRUE</code> .

## Details

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

## Value

A **list** of elements:

- `catalog`: The ID (small insertion and deletion) catalog with attributes added. See [as.catalog](#) for more details.
- `discarded.variants`: **Non-NULL only if** there are variants that were excluded from the analysis. See the added extra column `discarded.reason` for more details.
- `annotated.vcfs`: **Non-NULL only if** `return.annotated.vcfs = TRUE`. A list of data frames which contain the original VCF's ID mutation rows with three additional columns `seq.context.width`, `seq.context` and `ID.class` added. The category assignment of each ID mutation in VCF can be obtained from `ID.class` column.

**ID classification**

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 ID (small insertion and deletion) mutation classification.

See the documentation for [Canonicalize1Del](#) which 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 [FindDelMH](#)).

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

**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.s1.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,
    return.annotated.vcfs = FALSE,
    suppress.discarded.variants.warnings = TRUE
  )

```

## Arguments

<code>dir</code>	Pathname of the directory which contains <b>only</b> 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 region.
<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>	Deprecated. If there are ID variants whose REF do not match the extracted sequence from <code>ref.genome</code> , the function will automatically discard these variants and an element <code>discarded.variants</code> will appear in the return value. See <a href="#">AnnotateIDVCF</a> for more details.
<code>return.annotated.vcfs</code>	Logical. Whether to return the annotated VCFs with additional columns showing mutation class for each variant. Default is <code>FALSE</code> .
<code>suppress.discarded.variants.warnings</code>	Logical. Whether to suppress warning messages showing information about the discarded variants. Default is <code>TRUE</code> .

## Details

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

## Value

A **list** of elements:

- `catalog`: The ID (small insertion and deletion) catalog with attributes added. See [as.catalog](#) for more details.
- `discarded.variants`: **Non-NULL only if** there are variants that were excluded from the analysis. See the added extra column `discarded.reason` for more details.
- `annotated.vcfs`: **Non-NULL only if** `return.annotated.vcfs = TRUE`. A list of data frames which contain the original VCF's ID mutation rows with three additional columns `seq.context.width`, `seq.context` and `ID.class` added. The category assignment of each ID mutation in VCF can be obtained from `ID.class` column.

**ID classification**

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 ID (small insertion and deletion) mutation classification.

See the documentation for [Canonicalize1Del](#) which 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 [FindDelMH](#)).

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

**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. The function will find and merge adjacent SBS pairs into DBS if their VAFs are very similar. The default threshold value for VAF is 0.02.

**Usage**

```
StrelkaSBSVCFFilesToCatalog(
  files,
  ref.genome,
  trans.ranges = NULL,
  region = "unknown",
  names.of.VCFs = NULL,
  return.annotated.vcfs = FALSE,
```



```

    suppress.discarded.variants.warnings = TRUE
  )

```

## 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` Optional. If `ref.genome` specifies one of the [BSgenome](#) object

1. `BSgenome.Hsapiens.1000genomes.hs37d5`
2. `BSgenome.Hsapiens.UCSC.hg38`
3. `BSgenome.Mmusculus.UCSC.mm10`

then the function will infer `trans.ranges` automatically. Otherwise, user will need to provide the necessary `trans.ranges`. Please refer to [TranscriptRanges](#) for more details. If `is.null(trans.ranges)` do not add transcript range information.

`region` A character string designating a genomic region; see [as.catalog](#) and [ICAMS](#).

`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.

`return.annotated.vcfs` Logical. Whether to return the annotated VCFs with additional columns showing mutation class for each variant. Default is `FALSE`.

`suppress.discarded.variants.warnings` Logical. Whether to suppress warning messages showing information about the discarded variants. Default is `TRUE`.

## Details

This function calls [VCFsToSBSCatalogs](#) and [VCFsToDBSCatalogs](#).

## Value

A list containing the following objects:

- `catSBS96`, `catSBS192`, `catSBS1536`: Matrix of 3 SBS catalogs (one each for 96, 192, and 1536).
- `catDBS78`, `catDBS136`, `catDBS144`: Matrix of 3 DBS catalogs (one each for 78, 136, and 144).
- `discarded.variants`: **Non-NULL only if** there are variants that were excluded from the analysis. See the added extra column `discarded.reason` for more details.
- `annotated.vcfs`: **Non-NULL only if** `return.annotated.vcfs = TRUE`. A list of elements:
  - SBS: SBS VCF annotated by [AnnotateSBSVCF](#) with three new columns `SBS96.class`, `SBS192.class` and `SBS1536.class` showing the mutation class for each SBS variant.

- DBS: DBS VCF annotated by `AnnotateDBSVCF` with three new columns `DBS78.class`, `DBS136.class` and `DBS144.class` showing the mutation class for each DBS variant.

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.

### Note

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

### Comments

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

### Examples

```
file <- c(system.file("extdata/Strelka-SBS-vcf",
                    "Strelka.SBS.GRCh37.s1.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. The function will find and merge adjacent SBS pairs into DBS if their VAFs are very similar. The default threshold value for VAF is 0.02.

### Usage

```
StrelkaSBSVCFFilesToCatalogAndPlotToPdf(
  files,
  ref.genome,
  trans.ranges = NULL,
  region = "unknown",
  names.of.VCFs = NULL,
  output.file = "",
  return.annotated.vcfs = FALSE,
  suppress.discarded.variants.warnings = TRUE
)
```

**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>1. <code>BSgenome.Hsapiens.1000genomes.hs37d5</code></li> <li>2. <code>BSgenome.Hsapiens.UCSC.hg38</code></li> <li>3. <code>BSgenome.Mmusculus.UCSC.mm10</code></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>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>return.annotated.vcfs</code>	Logical. Whether to return the annotated VCFs with additional columns showing mutation class for each variant. Default is <code>FALSE</code> .
<code>suppress.discarded.variants.warnings</code>	Logical. Whether to suppress warning messages showing information about the discarded variants. Default is <code>TRUE</code> .

**Details**

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

**Value**

A list containing the following objects:

- `catSBS96`, `catSBS192`, `catSBS1536`: Matrix of 3 SBS catalogs (one each for 96, 192, and 1536).
- `catDBS78`, `catDBS136`, `catDBS144`: Matrix of 3 DBS catalogs (one each for 78, 136, and 144).
- `discarded.variants`: **Non-NULL only if** there are variants that were excluded from the analysis. See the added extra column `discarded.reason` for more details.
- `annotated.vcfs`: **Non-NULL only if** `return.annotated.vcfs = TRUE`. A list of elements:
  - SBS: SBS VCF annotated by [AnnotateSBSVCF](#) with three new columns `SBS96.class`, `SBS192.class` and `SBS1536.class` showing the mutation class for each SBS variant.

- DBS: DBS VCF annotated by `AnnotateDBSVCF` with three new columns `DBS78.class`, `DBS136.class` and `DBS144.class` showing the mutation class for each DBS variant.

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.

### Note

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

### Comments

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

### Examples

```
file <- c(system.file("extdata/Strelka-SBS-vcf",
                    "Strelka.SBS.GRCh37.s1.vcf",
                    package = "ICAMS"))
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  catalogs <-
    StrelkaSBSVCFFilesToCatalogAndPlotToPdf(file, ref.genome = "hg19",
                                             trans.ranges = trans.ranges.GRCh37,
                                             region = "genome",
                                             output.file =
                                             file.path(tempdir(), "StrelkaSBS"))}
```

---

## 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. The function will find and merge adjacent SBS pairs into DBS if their VAFs are very similar. The default threshold value for VAF is 0.02.

### Usage

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

```

    base.filename = "",
    return.annotated.vcfs = FALSE,
    suppress.discarded.variants.warnings = TRUE
)

```

### Arguments

<code>dir</code>	Pathname of the directory which contains <b>only</b> 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 region.
<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. <code>BSgenome.Hsapiens.1000genomes.hs37d5</code></li> <li>2. <code>BSgenome.Hsapiens.UCSC.hg38</code></li> <li>3. <code>BSgenome.Mmusculus.UCSC.mm10</code></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.
<code>return.annotated.vcfs</code>	Logical. Whether to return the annotated VCFs with additional columns showing mutation class for each variant. Default is <code>FALSE</code> .
<code>suppress.discarded.variants.warnings</code>	Logical. Whether to suppress warning messages showing information about the discarded variants. Default is <code>TRUE</code> .

### Details

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

### Value

A list containing the following objects:

- `catSBS96`, `catSBS192`, `catSBS1536`: Matrix of 3 SBS catalogs (one each for 96, 192, and 1536).

- `catDBS78`, `catDBS136`, `catDBS144`: Matrix of 3 DBS catalogs (one each for 78, 136, and 144).
- `discarded.variants`: **Non-NULL only if** there are variants that were excluded from the analysis. See the added extra column `discarded.reason` for more details.
- `annotated.vcfs`: **Non-NULL only if** `return.annotated.vcfs = TRUE`. A list of elements:
  - SBS: SBS VCF annotated by `AnnotateSBSVCF` with three new columns `SBS96.class`, `SBS192.class` and `SBS1536.class` showing the mutation class for each SBS variant.
  - DBS: DBS VCF annotated by `AnnotateDBSVCF` with three new columns `DBS78.class`, `DBS136.class` and `DBS144.class` showing the mutation class for each DBS variant.

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.

### Note

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

### Comments

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

### 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/genocode/Genocode\\_human/release\\_30/GRCh37\\_mapping/genocode.v30lift37.annotation.gff3.gz](ftp://ftp.ebi.ac.uk/pub/databases/genocode/Genocode_human/release_30/GRCh37_mapping/genocode.v30lift37.annotation.gff3.gz)

[ftp://ftp.ebi.ac.uk/pub/databases/genocode/Genocode\\_human/release\\_30/genocode.v30.annotation.gff3.gz](ftp://ftp.ebi.ac.uk/pub/databases/genocode/Genocode_human/release_30/genocode.v30.annotation.gff3.gz)

[ftp://ftp.ebi.ac.uk/pub/databases/genocode/Genocode\\_mouse/release\\_M21/genocode.vM21.annotation.gff3.gz](ftp://ftp.ebi.ac.uk/pub/databases/genocode/Genocode_mouse/release_M21/genocode.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](#).

## Rationale

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).

**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")}

```

---

VCFsToCatalogs

*Create SBS, DBS and Indel catalogs from VCFs*


---

**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**

```

VCFsToCatalogs(
  files,
  ref.genome,
  variant.caller = "unknown",
  num.of.cores = 1,
  trans.ranges = NULL,
  region = "unknown",
  names.of.VCFs = NULL,
  tumor.col.names = NA,
  filter.status = NULL,
  get.vaf.function = NULL,
  ...,
  max.vaf.diff = 0.02,
  return.annotated.vcfs = FALSE,
  suppress.discarded.variants.warnings = TRUE
)

```

**Arguments**

<code>files</code>	Character vector of file paths to the VCF files.
<code>ref.genome</code>	A <code>ref.genome</code> argument as described in <a href="#">ICAMS</a> .
<code>variant.caller</code>	Name of the variant caller that produces the VCF, can be either "strelka", "mutect", "freebayes" or "unknown". This information is needed to calculate the VAFs (variant allele frequencies). If variant caller is "unknown"(default) and

	<p><code>get.vaf.function</code> is NULL, then VAF and read depth will be NAs. If variant caller is "mutect", do <b>not</b> merge SBSs into DBS.</p>
<code>num.of.cores</code>	The number of cores to use. Not available on Windows unless <code>num.of.cores = 1</code> .
<code>trans.ranges</code>	<p>Optional. If <code>ref.genome</code> specifies one of the <a href="#">BSgenome</a> object</p> <ol style="list-style-type: none"> <li>1. <code>BSgenome.Hsapiens.1000genomes.hs37d5</code></li> <li>2. <code>BSgenome.Hsapiens.UCSC.hg38</code></li> <li>3. <code>BSgenome.Mmusculus.UCSC.mm10</code></li> </ol> <p>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.</p>
<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 NULL(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. Only applicable to <b>Mutect</b> VCFs. Character vector of column names in <b>Mutect</b> VCFs which contain the tumor sample information. The order of names in <code>tumor.col.names</code> should match the order of <b>Mutect</b> VCFs specified in <code>files</code> . If <code>tumor.col.names</code> is equal to NA(default), this function will use the 10th column in all the <b>Mutect</b> VCFs to calculate VAFs. See <a href="#">GetMutectVAF</a> for more details.
<code>filter.status</code>	The status indicating a variant has passed all filters. An example would be "PASS". Variants which don't have the specified <code>filter.status</code> in the FILTER column in VCF will be removed. If NULL(default), no variants will be removed from the original VCF.
<code>get.vaf.function</code>	Optional. Only applicable when <code>variant.caller</code> is " <b>unknown</b> ". Function to calculate VAF(variant allele frequency) and read depth information from original VCF. See <a href="#">GetMutectVAF</a> as an example. If NULL(default) and <code>variant.caller</code> is "unknown", then VAF and read depth will be NAs.
<code>...</code>	Optional arguments to <code>get.vaf.function</code> .
<code>max.vaf.diff</code>	<b>Not</b> applicable if <code>variant.caller = "mutect"</code> . The maximum difference of VAF, default value is 0.02. If the absolute difference of VAFs for adjacent SBSs is bigger than <code>max.vaf.diff</code> , then these adjacent SBSs are likely to be "merely" asynchronous single base mutations, opposed to a simultaneous doublet mutation or variants involving more than two consecutive bases.
<code>return.annotated.vcfs</code>	Logical. Whether to return the annotated VCFs with additional columns showing mutation class for each variant. Default is FALSE.
<code>suppress.discarded.variants.warnings</code>	Logical. Whether to suppress warning messages showing information about the discarded variants. Default is TRUE.

## Details

This function calls [VCFsToSBSCatalogs](#), [VCFsToDBSCatalogs](#) and [VCFsToIDCatalogs](#)

## Value

A list containing the following objects:

- `catSBS96`, `catSBS192`, `catSBS1536`: Matrix of 3 SBS catalogs (one each for 96, 192, and 1536).
- `catDBS78`, `catDBS136`, `catDBS144`: Matrix of 3 DBS catalogs (one each for 78, 136, and 144).
- `catID`: Matrix of ID (small insertion and deletion) catalog.
- `discarded.variants`: **Non-NULL only if** there are variants that were excluded from the analysis. See the added extra column `discarded.reason` for more details.
- `annotated.vcfs`: **Non-NULL only if** `return.annotated.vcfs = TRUE`. A list of elements:
  - SBS: SBS VCF annotated by [AnnotateSBSVCF](#) with three new columns `SBS96.class`, `SBS192.class` and `SBS1536.class` showing the mutation class for each SBS variant.
  - DBS: DBS VCF annotated by [AnnotateDBSVCF](#) with three new columns `DBS78.class`, `DBS136.class` and `DBS144.class` showing the mutation class for each DBS variant.
  - ID: ID VCF annotated by [AnnotateIDVCF](#) with one new column `ID.class` showing the mutation class for each ID variant.

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.

## ID classification

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 ID (small insertion and deletion) mutation classification.

See the documentation for [Canonicalize1Del](#) which 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 [FindDelMH](#)).

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

## 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+.

## Comments

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

**Examples**

```
file <- c(system.file("extdata/Mutect-vcf",
                    "Mutect.GRCh37.s1.vcf",
                    package = "ICAMS"))
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  catalogs <- VCFsToCatalogs(file, ref.genome = "hg19",
                             variant.caller = "mutect", region = "genome")}
```

---

VCFsToCatalogsAndPlotToPdf

*Create SBS, DBS and Indel catalogs from VCFs and plot them to PDF*

---

**Description**

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

**Usage**

```
VCFsToCatalogsAndPlotToPdf(
  files,
  output.dir,
  ref.genome,
  variant.caller = "unknown",
  num.of.cores = 1,
  trans.ranges = NULL,
  region = "unknown",
  names.of.VCFs = NULL,
  tumor.col.names = NA,
  filter.status = NULL,
  get.vaf.function = NULL,
  ...,
  max.vaf.diff = 0.02,
  base.filename = "",
  return.annotated.vcfs = FALSE,
  suppress.discarded.variants.warnings = TRUE
)
```

**Arguments**

<code>files</code>	Character vector of file paths to the VCF files.
<code>output.dir</code>	The directory where the PDF files will be saved.
<code>ref.genome</code>	A <code>ref.genome</code> argument as described in <a href="#">ICAMS</a> .
<code>variant.caller</code>	Name of the variant caller that produces the VCF, can be either "strelka", "mutect", "freebayes" or "unknown". This information is needed to calculate the VAFs (variant allele frequencies). If variant caller is "unknown"(default) and

	<p><code>get.vaf.function</code> is NULL, then VAF and read depth will be NAs. If variant caller is "mutect", do <b>not</b> merge SBSs into DBS.</p>
<code>num.of.cores</code>	The number of cores to use. Not available on Windows unless <code>num.of.cores = 1</code> .
<code>trans.ranges</code>	<p>Optional. If <code>ref.genome</code> specifies one of the <a href="#">BSgenome</a> object</p> <ol style="list-style-type: none"> <li>1. <code>BSgenome.Hsapiens.1000genomes.hs37d5</code></li> <li>2. <code>BSgenome.Hsapiens.UCSC.hg38</code></li> <li>3. <code>BSgenome.Mmusculus.UCSC.mm10</code></li> </ol> <p>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.</p>
<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 NULL (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. Only applicable to <b>Mutect</b> VCFs. Character vector of column names in <b>Mutect</b> VCFs which contain the tumor sample information. The order of names in <code>tumor.col.names</code> should match the order of <b>Mutect</b> VCFs specified in <code>files</code> . If <code>tumor.col.names</code> is equal to NA (default), this function will use the 10th column in all the <b>Mutect</b> VCFs to calculate VAFs. See <a href="#">GetMutectVAF</a> for more details.
<code>filter.status</code>	The status indicating a variant has passed all filters. An example would be "PASS". Variants which don't have the specified <code>filter.status</code> in the FILTER column in VCF will be removed. If NULL (default), no variants will be removed from the original VCF.
<code>get.vaf.function</code>	Optional. Only applicable when <code>variant.caller</code> is " <b>unknown</b> ". Function to calculate VAF (variant allele frequency) and read depth information from original VCF. See <a href="#">GetMutectVAF</a> as an example. If NULL (default) and <code>variant.caller</code> is "unknown", then VAF and read depth will be NAs.
<code>...</code>	Optional arguments to <code>get.vaf.function</code> .
<code>max.vaf.diff</code>	<b>Not</b> applicable if <code>variant.caller = "mutect"</code> . The maximum difference of VAF, default value is 0.02. If the absolute difference of VAFs for adjacent SBSs is bigger than <code>max.vaf.diff</code> , then these adjacent SBSs are likely to be "merely" asynchronous single base mutations, opposed to a simultaneous doublet mutation or variants involving more than two consecutive bases.
<code>base.filename</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>return.annotated.vcfs</code>	Logical. Whether to return the annotated VCFs with additional columns showing mutation class for each variant. Default is FALSE.

`suppress.discarded.variants.warnings`

Logical. Whether to suppress warning messages showing information about the discarded variants. Default is TRUE.

## Details

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

## Value

A list containing the following objects:

- `catSBS96`, `catSBS192`, `catSBS1536`: Matrix of 3 SBS catalogs (one each for 96, 192, and 1536).
- `catDBS78`, `catDBS136`, `catDBS144`: Matrix of 3 DBS catalogs (one each for 78, 136, and 144).
- `catID`: Matrix of ID (small insertion and deletion) catalog.
- `discarded.variants`: **Non-NULL only if** there are variants that were excluded from the analysis. See the added extra column `discarded.reason` for more details.
- `annotated.vcfs`: **Non-NULL only if** `return.annotated.vcfs = TRUE`. A list of elements:
  - SBS: SBS VCF annotated by [AnnotateSBSVCF](#) with three new columns `SBS96.class`, `SBS192.class` and `SBS1536.class` showing the mutation class for each SBS variant.
  - DBS: DBS VCF annotated by [AnnotateDBSVCF](#) with three new columns `DBS78.class`, `DBS136.class` and `DBS144.class` showing the mutation class for each DBS variant.
  - ID: ID VCF annotated by [AnnotateIDVCF](#) with one new column `ID.class` showing the mutation class for each ID variant.

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.

## ID classification

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 ID (small insertion and deletion) mutation classification.

See the documentation for [Canonicalize1Del](#) which 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 [FindDelMH](#)).

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

## 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+.

**Comments**

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

**Examples**

```
file <- c(system.file("extdata/Mutect-vcf",
                    "Mutect.GRCh37.s1.vcf",
                    package = "ICAMS"))
if (requireNamespace("BSgenome.Hsapiens.1000genomes.hs37d5", quietly = TRUE)) {
  catalogs <-
    VCFsToCatalogsAndPlotToPdf(file, ref.genome = "hg19",
                               output.dir = tempdir(),
                               variant.caller = "mutect",
                               region = "genome",
                               base.filename = "Mutect")}
```

---

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,
  num.of.cores = 1,
  trans.ranges = NULL,
  region = "unknown",
  return.annotated.vcfs = FALSE,
  suppress.discarded.variants.warnings = TRUE
)
```

**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>num.of.cores</code>	The number of cores to use. Not available on Windows unless <code>num.of.cores = 1</code> .
<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><code>BSgenome.Hsapiens.1000genomes.hs37d5</code></li> </ol>





---

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,
  num.of.cores = 1,
  region = "unknown",
  flag.mismatches = 0,
  return.annotated.vcfs = FALSE,
  suppress.discarded.variants.warnings = TRUE
)
```

### Arguments

<code>list.of.vcfs</code>	List of in-memory ID 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>num.of.cores</code>	The number of cores to use. Not available on Windows unless <code>num.of.cores = 1</code> .
<code>region</code>	A character string acting as a region identifier, one of "genome", "exome".
<code>flag.mismatches</code>	Deprecated. If there are ID variants whose REF do not match the extracted sequence from <code>ref.genome</code> , the function will automatically discard these variants and an element <code>discarded.variants</code> will appear in the return value. See <a href="#">AnnotateIDVCF</a> for more details.
<code>return.annotated.vcfs</code>	Logical. Whether to return the annotated VCFs with additional columns showing mutation class for each variant. Default is FALSE.
<code>suppress.discarded.variants.warnings</code>	Logical. Whether to suppress warning messages showing information about the discarded variants. Default is TRUE.

### Value

A **list** of elements:

- `catalog`: The ID (small insertion and deletion) catalog with attributes added. See [as.catalog](#) for details.

- `discarded.variants`: **Non-NULL only if** there are variants that were excluded from the analysis. See the added extra column `discarded.reason` for more details.
- `annotated.vcfs`: **Non-NULL only if** `return.annotated.vcfs = TRUE`. A list of data frames which contain the original VCF's ID mutation rows with three additional columns `seq.context.width`, `seq.context` and `ID.class` added. The category assignment of each ID mutation in VCF can be obtained from `ID.class` column.

### 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+.

### ID classification

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 ID (small insertion and deletion) mutation classification.

See the documentation for [Canonicalize1Del](#) which 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 [FindDelMH](#)).

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

### Examples

```
file <- c(system.file("extdata/Strelka-ID-vcf/",
                    "Strelka.ID.GRCh37.s1.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,
  num.of.cores = 1,
  trans.ranges = NULL,
  region = "unknown",
  return.annotated.vcfs = FALSE,
  suppress.discarded.variants.warnings = TRUE
)
```

**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>num.of.cores</code>	The number of cores to use. Not available on Windows unless <code>num.of.cores = 1</code> .
<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><code>BSgenome.Hsapiens.1000genomes.hs37d5</code></li> <li><code>BSgenome.Hsapiens.UCSC.hg38</code></li> <li><code>BSgenome.Mmusculus.UCSC.mm10</code></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>return.annotated.vcfs</code>	Logical. Whether to return the annotated VCFs with additional columns showing mutation class for each variant. Default is <code>FALSE</code> .
<code>suppress.discarded.variants.warnings</code>	Logical. Whether to suppress warning messages showing information about the discarded variants. Default is <code>TRUE</code> .

**Value**

A list containing the following objects:

- `catSBS96`, `catSBS192`, `catSBS1536`: Matrix of 3 SBS catalogs (one each for 96, 192, and 1536).
- `discarded.variants`: **Non-NULL only if** there are variants that were excluded from the analysis. See the added extra column `discarded.reason` for more details.
- `annotated.vcfs`: **Non-NULL only if** `return.annotated.vcfs = TRUE`. SBS VCF annotated by [AnnotateSBSVCF](#) with three new columns `SBS96.class`, `SBS192.class` and `SBS1536.class` showing the mutation class for each SBS variant.

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.s1.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")}
```

---

VCFsToZipFile

*Create a zip file which contains catalogs and plot PDFs from VCFs*

---

### Description

Create 3 SBS catalogs (96, 192, 1536), 3 DBS catalogs (78, 136, 144) and Indel catalog from the 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

```
VCFsToZipFile(
  dir,
  files,
  zipfile,
  ref.genome,
  variant.caller = "unknown",
  num.of.cores = 1,
  trans.ranges = NULL,
  region = "unknown",
  names.of.VCFs = NULL,
  tumor.col.names = NA,
  filter.status = NULL,
  get.vaf.function = NULL,
  ...,
```

```

max.vaf.diff = 0.02,
base.filename = "",
return.annotated.vcfs = FALSE,
suppress.discarded.variants.warnings = TRUE
)

```

## Arguments

<code>dir</code>	Pathname of the directory which contains VCFs that come from the <b>same</b> variant caller. Each VCF <b>must</b> have a file extension ".vcf" (case insensitive) and share the <b>same</b> ref.genome and region.
<code>files</code>	Character vector of file paths to the VCF files. Only <b>one</b> of argument <code>dir</code> or <code>files</code> need to be specified.
<code>zipfile</code>	Pathname of the zip file to be created.
<code>ref.genome</code>	A ref.genome argument as described in <a href="#">ICAMS</a> .
<code>variant.caller</code>	Name of the variant caller that produces the VCF, can be either "strelka", "mutect", "freebayes" or "unknown". This information is needed to calculate the VAFs (variant allele frequencies). If variant caller is "unknown"(default) and <code>get.vaf</code> function is NULL, then VAF and read depth will be NAs. If variant caller is "mutect", do <b>not</b> merge SBSs into DBS.
<code>num.of.cores</code>	The number of cores to use. Not available on Windows unless <code>num.of.cores = 1</code> .
<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>BSgenome.Hsapiens.1000genomes.hs37d5</li> <li>BSgenome.Hsapiens.UCSC.hg38</li> <li>BSgenome.Mmusculus.UCSC.mm10</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 NULL(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. Only applicable to <b>Mutect</b> VCFs. Character vector of column names in <b>Mutect</b> VCFs which contain the tumor sample information. The order of names in <code>tumor.col.names</code> should match the order of <b>Mutect</b> VCFs specified in <code>files</code> . If <code>tumor.col.names</code> is equal to NA(default), this function will use the 10th column in all the <b>Mutect</b> VCFs to calculate VAFs. See <a href="#">GetMutectVAF</a> for more details.
<code>filter.status</code>	The status indicating a variant has passed all filters. An example would be "PASS". Variants which don't have the specified <code>filter.status</code> in the FILTER

	column in VCF will be removed. If NULL(default), no variants will be removed from the original VCF.
<code>get.vaf.function</code>	Optional. Only applicable when <code>variant.caller</code> is <b>"unknown"</b> . Function to calculate VAF(variant allele frequency) and read depth information from original VCF. See <a href="#">GetMutectVAF</a> as an example. If NULL(default) and <code>variant.caller</code> is "unknown", then VAF and read depth will be NAs.
<code>...</code>	Optional arguments to <code>get.vaf.function</code> .
<code>max.vaf.diff</code>	<b>Not</b> applicable if <code>variant.caller = "mutect"</code> . The maximum difference of VAF, default value is 0.02. If the absolute difference of VAFs for adjacent SBSs is bigger than <code>max.vaf.diff</code> , then these adjacent SBSs are likely to be "merely" asynchronous single base mutations, opposed to a simultaneous doublet mutation or variants involving more than two consecutive bases.
<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>return.annotated.vcfs</code>	Logical. Whether to return the annotated VCFs with additional columns showing mutation class for each variant. Default is FALSE.
<code>suppress.discarded.variants.warnings</code>	Logical. Whether to suppress warning messages showing information about the discarded variants. Default is TRUE.

## Details

This function calls [VCFsToCatalogs](#), [PlotCatalogToPdf](#), [WriteCatalog](#) and `zip::zipr`.

## Value

A list containing the following objects:

- `catSBS96`, `catSBS192`, `catSBS1536`: Matrix of 3 SBS catalogs (one each for 96, 192, and 1536).
- `catDBS78`, `catDBS136`, `catDBS144`: Matrix of 3 DBS catalogs (one each for 78, 136, and 144).
- `catID`: Matrix of ID (small insertion and deletion) catalog.
- `discarded.variants`: **Non-NULL only if** there are variants that were excluded from the analysis. See the added extra column `discarded.reason` for more details.
- `annotated.vcfs`: **Non-NULL only if** `return.annotated.vcfs = TRUE`. A list of elements:
  - SBS: SBS VCF annotated by [AnnotateSBSVCF](#) with three new columns `SBS96.class`, `SBS192.class` and `SBS1536.class` showing the mutation class for each SBS variant.
  - DBS: DBS VCF annotated by [AnnotateDBSVCF](#) with three new columns `DBS78.class`, `DBS136.class` and `DBS144.class` showing the mutation class for each DBS variant.
  - ID: ID VCF annotated by [AnnotateIDVCF](#) with one new column `ID.class` showing the mutation class for each ID variant.

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.

**ID classification**

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 ID (small insertion and deletion) mutation classification.

See the documentation for `Canonicalize1Del` which 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 `FindDelMH`).

See the code for unexported function `CanonicalizeID` and the functions it calls for handling of insertions.

**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+.

**Comments**

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

**Examples**

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

---

WriteCatalog

*Write a catalog*

---

**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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