

# Package ‘VariantFiltering’

September 4, 2025

**Type** Package

**Title** Filtering of coding and non-coding genetic variants

**Description** Filter genetic variants using different criteria such as inheritance model, amino acid change consequence, minor allele frequencies across human populations, splice site strength, conservation, etc.

**Version** 1.44.0

**License** Artistic-2.0

**Depends** R (>= 3.5.0), methods, BiocGenerics (>= 0.25.1), VariantAnnotation (>= 1.13.29)

**Imports** utils, stats, Biobase, S4Vectors (>= 0.9.25), IRanges (>= 2.3.23), RBGL, graph, AnnotationDbi, BiocParallel, Biostrings (>= 2.33.11), GenomeInfoDb (>= 1.3.6), GenomicRanges (>= 1.19.13), SummarizedExperiment, GenomicFeatures, Rsamtools (>= 1.17.8), BSgenome, GenomicScores (>= 1.0.0), Gviz, shiny, shinythemes, shinyjs, DT, shinyTree

**Suggests** RUnit, BiocStyle, org.Hs.eg.db, BSgenome.Hsapiens.1000genomes.hs37d5, TxDb.Hsapiens.UCSC.hg19.knownGene, SNPlocs.Hsapiens.dbSNP144.GRCh37, MafDb.1Kgenomes.phase1.hs37d5, phastCons100way.UCSC.hg19, PolyPhen.Hsapiens.dbSNP131, SIFT.Hsapiens.dbSNP137

**LinkingTo** S4Vectors, IRanges, XVector, Biostrings

**URL** <https://github.com/rcastelo/VariantFiltering>

**BugReports** <https://github.com/rcastelo/VariantFiltering/issues>

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VariantFiltering-package  
*Filtering of coding and non-coding genetic variants*

---

Description

The VariantFiltering package filters coding and non-coding genetic variants using different criteria such as an inheritance model (autosomal recessive -both, homozygous and heterozygous-, autosomal dominant, X-linked and *de novo*), amino acid change consequence, minor allele frequencies, cryptic splice site potential, conservation, etc.

Functions

- [autosomalRecessiveHomozygous](#) identify homozygous variants in the affected individual(s) while the unaffected ones present these same variants but in heterozygous state. Autosomal recessive inheritance pattern.

- [autosomalRecessiveHeterozygous](#) identify variants grouped by genes with two (or more) heterogeneous alleles (at least one on each allele, i.e. coming from each parent). Autosomal recessive inheritance pattern.
- [autosomalDominant](#) identify variants present in all the affected individual(s) discarding the ones that also occur in at least one of the unaffected subjects. Autosomal dominant inheritance pattern.
- [xLinked](#) identify variants that appear only in the X chromosome of the unaffected females as heterozygous, don't appear in the unaffected males analyzed and finally are present (as homozygous) in the affected male(s). X-linked inheritance pattern.
- [deNovo](#) identify variants in the affected individual that have not been inherited.
- [unrelatedIndividuals](#) annotate variants without filtering by any inheritance pattern.

### Author(s)

Dei M. Elurbe and Robert Castelo.

Maintainer: Robert Castelo <robert.castelo@upf.edu>

### References

Elurbe D.M., Mila, M., Castelo, R. VariantFiltering: filtering of coding and non-coding genetic variants, in preparation.

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|                   |                                                |
|-------------------|------------------------------------------------|
| autosomalDominant | <i>Autosomal dominant inheritance analysis</i> |
|-------------------|------------------------------------------------|

---

### Description

This function identifies variants present in all the affected individual(s) discarding the ones that also occur in at least one of the unaffected subjects.

### Usage

```
## S4 method for signature 'VariantFilteringParam'
autosomalDominant(param,
                   svparam=ScanVcfParam(),
                   use=c("everything", "complete.obs", "all.obs"),
                   BPPARAM=bpparam("SerialParam"))
```

### Arguments

|       |                                                                                                                                                                                                                                                                                                           |
|-------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| param | A <a href="#">VariantFilteringParam</a> object built from a multisample VCF file with at least one affected individual and zero or more unaffected ones, and from a PED file specifying the family relationships among individuals as well as their gender and phenotype status (affected or unaffected). |
|-------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|

|         |                                                                                                                                                                                                                                                                                                                                                                                                                   |
|---------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| svparam | An instance of a ScanVcfParam object to enable analyzing a subset of variants and samples. This object is passed internally to a call to the <code>readVcf()</code> function in the VariantAnnotation package, see its help page for a complete description of this functionality.                                                                                                                                |
| use     | character string specifying the policy to apply on missing genotypes when comparing them. This policy can be either "everything" (default), "complete.obs" or "all.obs". The default policy ("everything") will propagate NA truth values using the behavior of the R logical operators, with the exception that when the final truth value associated with a variant is NA, the variant is ultimately discarded. |
| BPPARAM | An object of class <code>BiocParallelParam</code> specifying parameters related to the parallel execution of some of the tasks and calculations within this function. See function <code>bpparam()</code> from the BiocParallel package.                                                                                                                                                                          |

### Details

This function requires as an input a `VariantFilteringParam` class object built from an input multisample VCF file, along with a PED file.

### Value

An object of class `VariantFilteringResults` including functional annotations on variants segregating according to an autosomal dominant inheritance model.

### Author(s)

Dei M. Elurbe and R. Castelo

### References

Elurbe D.M., Mila, M., Castelo, R. The VariantFiltering package, in preparation.

### See Also

[autosomalRecessiveHomozygous](#) [autosomalRecessiveHeterozygous](#) [xLinked](#) [deNovo](#) [unrelatedIndividuals](#) [VariantFilteringResults](#)

### Examples

```
## Not run:

CEUvcf <- file.path(system.file("extdata", package="VariantFiltering"), "CEUtrio.vcf.bgz")
CEUped <- file.path(system.file("extdata", package="VariantFiltering"), "CEUtrio.ped")
param <- VariantFilteringParam(vcfFilename=CEUvcf,
                              pedFilename=CEUped)
aDo <- autosomalDominant(param)
aDo

## End(Not run)
```

---

`autosomalRecessiveHeterozygous`*Autosomal recessive inheritance analysis: Heterozygous*

---

## Description

This function aims to analyze the variants of the unaffected individuals, storing and grouping the heterozygous ones by gene. The affected individuals ought present two or more different heterozygous changes in the gene, and at least one of them shall come from each parent.

## Usage

```
## S4 method for signature 'VariantFilteringParam'
autosomalRecessiveHeterozygous(param,
                                svparam=ScanVcfParam(),
                                BPPARAM=bpparam("SerialParam"))
```

## Arguments

|                      |                                                                                                                                                                                                                                                                                                                       |
|----------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <code>param</code>   | A <a href="#">VariantFilteringParam</a> object built from a multisample VCF file with at least one affected individual and two or more unaffected ones, and from a PED file specyfing the family relationships among individuals as well as their gender and phenotype status (affected or unaffected).               |
| <code>svparam</code> | An instance of a <a href="#">ScanVcfParam</a> object to enable analyzing a subset of variants and samples. This object is passed internally to a call to the <a href="#">readVcf()</a> function in the <a href="#">VariantAnnotation</a> package, see its help page for a complete description of this functionality. |
| <code>BPPARAM</code> | An object of class <a href="#">BiocParallelParam</a> specifying parameters related to the parallel execution of some of the tasks and calculations within this function. See function <a href="#">bpparam()</a> from the <a href="#">BiocParallel</a> package.                                                        |

## Details

This function requires as an input a [VariantFilteringParam](#) class object built from an input multisample VCF file, along with a PED file.

## Value

An object of class [VariantFilteringResults](#) including functional annotations on all selected variants.

## Author(s)

Dei M. Elurbe and R. Castelo

## References

Elurbe D.M., Mila, M., Castelo, R. The [VariantFiltering](#) package, in preparation.

**See Also**

[autosomalRecessiveHomozygous autosomalDominant xLinked deNovo VariantFilteringResults](#)

**Examples**

```
## Not run:

CEUvcf <- file.path(system.file("extdata", package="VariantFiltering"), "CEUtrio.vcf.bgz")
CEUped <- file.path(system.file("extdata", package="VariantFiltering"), "CEUtrio.ped")
param <- VariantFilteringParam(vcfFilename=CEUvcf,
                              pedFilename=CEUped)
reHet <- autosomalRecessiveHeterozygous(param)
reHet

## End(Not run)
```

---

autosomalRecessiveHomozygous

*Autosomal recessive inheritance analysis: Homozygous*

---

**Description**

This function works analyzing the variants of the unaffected individuals storing the common heterozygous ones and comparing them with the common homozygous variants between the affected subjects.

**Usage**

```
## S4 method for signature 'VariantFilteringParam'
autosomalRecessiveHomozygous(param,
                              svparam=ScanVcfParam(),
                              use=c("everything", "complete.obs", "all.obs"),
                              includeHomRef=FALSE,
                              age.of.onset,
                              phenocopies,
                              BPPARAM=bpparam("SerialParam"))
```

**Arguments**

|         |                                                                                                                                                                                                                                                                                                           |
|---------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| param   | A <a href="#">VariantFilteringParam</a> object built from a multisample VCF file with at least one affected individual and zero or more unaffected ones, and from a PED file specifying the family relationships among individuals as well as their gender and phenotype status (affected or unaffected). |
| svparam | An instance of a ScanVcfParam object to enable analyzing a subset of variants and samples. This object is passed internally to a call to the <a href="#">readVcf()</a> function in the VariantAnnotation package, see its help page for a complete description of this functionality.                     |

|               |                                                                                                                                                                                                                                                                                                                                                                                                                   |
|---------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| use           | character string specifying the policy to apply on missing genotypes when comparing them. This policy can be either "everything" (default), "complete.obs" or "all.obs". The default policy ("everything") will propagate NA truth values using the behavior of the R logical operators, with the exception that when the final truth value associated with a variant is NA, the variant is ultimately discarded. |
| includeHomRef | logical value specifying whether the homozygous reference genotype determines the affected phenotype (TRUE) or not (FALSE, default).                                                                                                                                                                                                                                                                              |
| age.of.onset  | numerical value specifying the age of onset at which individuals below that age are set their phenotype to unknown. This argument is currently experimental.                                                                                                                                                                                                                                                      |
| phenocopies   | numerical value specifying the maximum fraction of affected individuals not sharing the homozygous genotype. This argument is currently experimental                                                                                                                                                                                                                                                              |
| BPPARAM       | An VariantFilteringParam object containing VCF file(s). From 1 to 5 independent files for affected individuals and 0 to 5 more for unaffected ones (up to 10 individuals). If the VCF is a multi-sample, it can contain the same amount of individuals divided likewise.                                                                                                                                          |

### Details

This function requires as an input a [VariantFilteringParam](#) class object built from an input multi-sample VCF file, along with a PED file.

### Value

An object of class [VariantFilteringResults](#) including functional annotations on all selected variants.

### Author(s)

Dei M. Elurbe and R. Castelo

### References

Elurbe D.M., Mila, M., Castelo, R. The VariantFiltering package, in preparation.

### See Also

[autosomalRecessiveHeterozygous](#) [autosomalDominant](#) [xLinked](#) [deNovo](#) [unrelatedIndividuals](#) [VariantFilteringResults](#)

### Examples

```
## Not run:
library(VariantFiltering)

CEUvcf <- file.path(system.file("extdata", package="VariantFiltering"),
                    "CEUtrio.vcf.bgz")
CEUped <- file.path(system.file("extdata", package="VariantFiltering"),
                    "CEUtrio.ped")
```

```

param <- VariantFilteringParam(vcfFilename=CEUvcf, pedFilename=CEUped)
reHo <- autosomalRecessiveHomozygous(param)
reHo

## End(Not run)

```

deNovo

*De Novo variants analysis*

## Description

This function has been created in order to search and annotate *de novo* variants in one individual, discarding the ones shared with his/her parents.

## Usage

```

## S4 method for signature 'VariantFilteringParam'
deNovo(param,
        svparam=ScanVcfParam(),
        use=c("everything", "complete.obs", "all.obs"),
        BPPARAM=bpparam("SerialParam"))

```

## Arguments

|         |                                                                                                                                                                                                                                                                                                                                                                                                                   |
|---------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| param   | A <a href="#">VariantFilteringParam</a> object built from a multisample VCF file with at least one affected individual and zero or more unaffected ones, and from a PED file specifying the family relationships among individuals as well as their gender and phenotype status (affected or unaffected).                                                                                                         |
| svparam | An instance of a <a href="#">ScanVcfParam</a> object to enable analyzing a subset of variants and samples. This object is passed internally to a call to the <a href="#">readVcf()</a> function in the <a href="#">VariantAnnotation</a> package, see its help page for a complete description of this functionality.                                                                                             |
| use     | character string specifying the policy to apply on missing genotypes when comparing them. This policy can be either "everything" (default), "complete.obs" or "all.obs". The default policy ("everything") will propagate NA truth values using the behavior of the R logical operators, with the exception that when the final truth value associated with a variant is NA, the variant is ultimately discarded. |
| BPPARAM | An object of class <a href="#">BiocParallelParam</a> specifying parameters related to the parallel execution of some of the tasks and calculations within this function. See function <a href="#">bpparam()</a> from the <a href="#">BiocParallel</a> package.                                                                                                                                                    |

## Details

This function requires as an input a [VariantFilteringParam](#) class object built from an input multisample VCF file, along with a PED file.

**Value**

An object of class [VariantFilteringResults](#) including functional annotations on all selected variants.

**Author(s)**

Dei M. Elurbe and R. Castelo

**References**

Elurbe D.M., Mila, M., Castelo, R. The VariantFiltering package, in preparation.

**See Also**

[autosomalRecessiveHomozygous](#) [autosomalDominant](#) [autosomalRecessiveHeterozygous](#) [xLinked](#) [VariantFilteringResults](#)

**Examples**

```
## Not run:

CEUvcf <- file.path(system.file("extdata", package="VariantFiltering"), "CEUtrio.vcf.bgz")
CEUped <- file.path(system.file("extdata", package="VariantFiltering"), "CEUtrio.ped")
param <- VariantFilteringParam(vcfFilename=CEUvcf,
                              pedFilename=CEUped)
deNo <- deNovo(param)
deNo

## End(Not run)
```

---

GenePhylostrataDb-class

*PhastConsDb class*

---

**Description**

Class for storing gene-level conservation information in the form of levels of phylogenetic strata; see Neme and Tautz (2013).

**Usage**

```
## S4 method for signature 'GenePhylostrataDb'
genePhylostrata(object)
## S4 method for signature 'GenePhylostrataDb'
genePhylostratum(object, ids)
## S4 method for signature 'GenePhylostrataDb'
organism(object)
```

**Arguments**

|        |                                                                                                                                                                                       |
|--------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| object | A GenePhylostrataDb object.                                                                                                                                                           |
| ids    | A string character vector with the gene identifiers to fetch their phylostrata. These identifiers can be only either Ensembl Gene Identifiers (ENSGXXXXX) or Entrez Gene Identifiers. |

**Details**

The GenePhylostrataDb class and associated methods serve the purpose of storing and manipulating gene-level conservation information in the form of levels of phylogenetic strata (Neme and Tautz, 2013). One such objects is created at loading time by the VariantFiltering package with the constructor function GenePhylostrataDb(), and it is called humanGenesPhylostrata.

**Value**

.

**Author(s)**

R. Castelo

**Source**

<http://genomebiology.com/content/supplementary/1471-2164-14-117-s1.xlsx>

**References**

Neme, R. and Tautz, D. Phylogenetic patterns of emergence of new genes support a model of frequent de novo evolution. BMC Genomics, 14:117, 2013

**See Also**

[phastCons100way.UCSC.hg19](#)

**Examples**

```
humanGenesPhylostrata
```

---

```
readAARadicalChangeMatrix
```

*Read matrix of amino acid radical changes*

---

**Description**

Function to read and parse a tab-separated file of amino acid properties into a matrix of logical values indicating whether the change of one amino acid by another can be considered radical or conservative according to the chemical properties specified in the input file.

**Usage**

```
readAARadicalChangeMatrix(file)
```

**Arguments**

|             |                                                                                                                                                                                                                                                                                                                                                    |
|-------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <b>file</b> | A file containing a classification of amino acids with respect to one or more chemical properties. Its particular format should match the one from the file called <code>AA_chemical_properties_HanadaGojoboriLi2006.tsv</code> found in the <code>extdata</code> folder of this package. This file is based on Table 1 from Hanada et al. (2006). |
|-------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|

**Details**

The input file should contain one or more columns each of them forming a logical mask specifying sets of amino acids sharing some chemical property.

**Value**

An squared symmetric matrix with as many rows and columns as amino acids and whose cells contain logical values. These values are set to `TRUE` whenever the amino acid change implied by row and column is considered radical and `FALSE` when considered conservative. Amino acid changes within a chemical property are defined as conservative and radical otherwise.

**Author(s)**

R. Castelo

**References**

Hanada, K., Gojobori, T. and Li, W. Radical amino acid change versus positive selection in the evolution of viral envelope proteins. *Gene*, 385:83-88, 2006.

**See Also**

[VariantFilteringParam](#)

**Examples**

```
aamat <- readAARadicalChangeMatrix(file.path(system.file("extdata", package="VariantFiltering"),
                                                         "AA_chemical_properties_HanadaGojoboriLi2006.tsv"))
aamat[1:5, 1:5]
```

---

unrelatedIndividuals    *Analysis designed to be applied over a group of unrelated individuals*

---

## Description

This function is designed to create an object to deepen into the variants presented by a group of unrelated individuals

## Usage

```
## S4 method for signature 'VariantFilteringParam'
unrelatedIndividuals(param,
                     svparam=ScanVcfParam(),
                     BPPARAM=bpparam("SerialParam"))
```

## Arguments

|         |                                                                                                                                                                                                                                                                                                                       |
|---------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| param   | A <a href="#">VariantFilteringParam</a> object built from a multisample VCF file.                                                                                                                                                                                                                                     |
| svparam | An instance of a <a href="#">ScanVcfParam</a> object to enable analyzing a subset of variants and samples. This object is passed internally to a call to the <a href="#">readVcf()</a> function in the <a href="#">VariantAnnotation</a> package, see its help page for a complete description of this functionality. |
| BPPARAM | An object of class <a href="#">BiocParallelParam</a> specifying parameters related to the parallel execution of some of the tasks and calculations within this function. See function <a href="#">bpparam()</a> from the <a href="#">BiocParallel</a> package.                                                        |

## Details

This function requires as an input a [VariantFilteringParam](#) class object built from an input multisample VCF file.

## Value

An object of class [VariantFilteringResults](#) including functional annotations on all variants.

## Author(s)

Dei M. Elurbe and R. Castelo

## References

Elurbe D.M., Mila, M., Castelo, R. The VariantFiltering package, in preparation.

## See Also

[autosomalRecessiveHomozygous](#) [autosomalRecessiveHeterozygous](#) [autosomalDominant](#) [deNovo](#)  
[xLinked](#) [VariantFilteringResults](#)

**Examples**

```
## Not run:

CEUvcf <- file.path(system.file("extdata", package="VariantFiltering"), "CEUtrio.vcf.bgz")
param <- VariantFilteringParam(vcfFilename=CEUvcf)
uInd <- unrelatedIndividuals(param)
uInd

## End(Not run)
```

---

VariantFiltering-defunct

*Defunct functions in package 'VariantFiltering'*

---

**Description**

There are currently no defunct functions in 'VariantFiltering'.

**Author(s)**

Robert Castelo <robert.castelo@upf.edu>

---

VariantFiltering-deprecated

*Deprecated functions in package 'VariantFiltering'*

---

**Description**

There are currently no deprecated functions in 'VariantFiltering'.

**Details**

There are currently no deprecated functions.

**Author(s)**

Robert Castelo <robert.castelo@upf.edu>

---

VariantFilteringParam-class

*VariantFiltering parameter class*


---

## Description

The class VariantFilteringParam is defined to ease configuring the call to the functions that filter input genetic variants according to a desired segregating inheritance model (xLinked(), autosomalRecessiveHomozygous(), etc).

## Usage

```
VariantFilteringParam(vcfFilename, pedFilename=NA_character_,
                      bsgenome="BSgenome.Hsapiens.1000genomes.hs37d5",
                      orgdb="org.Hs.eg.db",
                      txdb="TxDb.Hsapiens.UCSC.hg19.knownGene",
                      snpdb="SNPlocs.Hsapiens.dbSNP144.GRCh37",
                      weightMatricesFileNames=NA,
                      weightMatricesLocations=rep(list(variantLocations()), length(weightMatricesFileNames)),
                      weightMatricesStrictLocations=rep(list(FALSE), length(weightMatricesFileNames)),
                      radicalAAchangeFilename=file.path(system.file("extdata",
                                                                    package="VariantFiltering"),
                                                         "AA_chemical_properties_HanadaGojoboriLi2006.tsv"),
                      codonusageFilename=file.path(system.file("extdata",
                                                                package="VariantFiltering"),
                                                     "humanCodonUsage.txt"),
                      geneticCode=getGeneticCode("SGC0"),
                      allTranscripts=FALSE,
                      regionAnnotations=list(CodingVariants(), IntronVariants(),
                                             FiveSpliceSiteVariants(), ThreeSpliceSiteVariants(),
                                             PromoterVariants(), FiveUTRVariants(), ThreeUTRVariants()),
                      intergenic=FALSE,
                      otherAnnotations=c("MafDb.1Kgenomes.phase1.hs37d5",
                                         "PolyPhen.Hsapiens.dbSNP131",
                                         "SIFT.Hsapiens.dbSNP137",
                                         "phastCons100way.UCSC.hg19",
                                         "humanGenesPhylostrata"),
                      geneKeytype=NA_character_,
                      yieldSize=NA_integer_)
## S4 method for signature 'VariantFilteringParam'
show(object)
## S4 method for signature 'VariantFilteringParam'
x$name
## S4 method for signature 'VariantFilteringParam'
names(x)
```

**Arguments**

|                                            |                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 |
|--------------------------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <code>vcfFilename</code>                   | Character string of the input VCF file name.                                                                                                                                                                                                                                                                                                                                                                                                                                    |
| <code>pedFilename</code>                   | Character string of the pedigree file name in PED format.                                                                                                                                                                                                                                                                                                                                                                                                                       |
| <code>bsgenome</code>                      | Character string of a genome annotation package ( <code>BSgenome.Hsapiens.1000genomes.hs37d5</code> by default).                                                                                                                                                                                                                                                                                                                                                                |
| <code>orgdb</code>                         | Character string of a gene-centric annotation package ( <code>org.Hs.eg.db</code> by default).                                                                                                                                                                                                                                                                                                                                                                                  |
| <code>txdb</code>                          | Character string of a transcript-centric annotation package ( <code>TxDb.Hsapiens.UCSC.hg19.knownGene</code> by default). The package <code>GenomicFeatures</code> provides infrastructure to build such annotation packages from different sources such as online UCSC tracks, Biomart tables, or GFF files.                                                                                                                                                                   |
| <code>snpdb</code>                         | Character string of a SNP-centric annotation package ( <code>SNPlocs.Hsapiens.dbSNP.20120608</code> by default).                                                                                                                                                                                                                                                                                                                                                                |
| <code>weightMatricesFileNames</code>       | Character string of filenames of position weight matrices for binding site recognition. The default NA value indicates that no binding sites will be scored. To use this feature to score, for instance, splice sites in human, assign to this argument the function <code>spliceSiteMatricesHuman()</code> . See the files ( <code>hsap.donors.hcmc10_15_1.ibn</code> and <code>hsap.acceptors.hcmc10_15_1.ibn</code> ) returned by this function for details on their format. |
| <code>weightMatricesLocations</code>       | Keywords of the annotated locations to variants under which a weight matrix will be used for scoring binding sites. This argument is only used when <code>weightMatricesFileNames!=NA</code> and, in such case, then more than one matrix is provided, this argument should be a list of character vectors with as many elements as matrices given in <code>weightMatricesFileNames</code> . The possible values can be obtained by typing <code>variantLocations()</code> .    |
| <code>weightMatricesStrictLocations</code> | Logical vector flagging whether a weight matrix should be scoring binding sites strictly within the boundaries of the given locations. This argument is only used when <code>weightMatricesFileNames!=NA</code> and, in such case, then more than one matrix is provided, this argument should be a list of logical vectors with as many elements as matrices given in <code>weightMatricesFileNames</code> .                                                                   |
| <code>radicalAAchangeFilename</code>       | Name of a tab-separated text file containing chemical properties of amino acids. These properties are interpreted such that amino acid changes within a property are considered "conservative" and between properties are considered "radical". See the default file ( <code>AA_chemical_properties_HanadaGojoboriLi2006.tsv</code> ) for details on its format.                                                                                                                |
| <code>codonusageFilename</code>            | Name of a text file containing the codon usage.                                                                                                                                                                                                                                                                                                                                                                                                                                 |
| <code>geneticCode</code>                   | Name character vector of length 64 describing the genetic code. The default value is <code>getGeneticCode("SGC0")</code> , the standard genetic code. An alternative genetic code, for instance, is <code>getGeneticCode("SGC1")</code> , the vertebrate mitochondrial genetic code. See <a href="#">getGeneticCode</a> in the <code>Biostrings</code> package for further details.                                                                                             |

|                                |                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              |
|--------------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <code>allTranscripts</code>    | Logical. This option allows the user to choose between working with all the transcripts affected by the variant ( <code>allTranscripts=TRUE</code> ) or with only one transcript per variant.                                                                                                                                                                                                                                                                                                                                                                                |
| <code>regionAnnotations</code> | List of <a href="#">VariantType-class</a> objects defining what regions to annotate.                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         |
| <code>intergenic</code>        | Logical. When <code>TRUE</code> , the intergenic variants are also annotated.                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                |
| <code>otherAnnotations</code>  | Character vector of names of annotation packages or annotation objects.                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      |
| <code>geneKeytype</code>       | Character vector of the type of key gene identifier provided by the transcript-centric (TxDb) annotation package to interrogate the organism-centric (OrgDb) annotation package. The default value ( <code>NA_character_</code> ) indicates that it will be assumed to be an Entrez identifier unless the values in the GENEID column returned by the TxDb package start with ENSG and then it will be assumed that they are Ensembl gene identifiers, or with one of NM_, NP_, NR_, XM_, XP_, XR_ or YP_ and then it will be assumed that they are RefSeq gene identifiers. |
| <code>yieldSize</code>         | Number of variants to yield each time the input VCF file is read. This argument is passed to the <a href="#">TabixFile</a> function when opening the input VCF file and it allows to iterate through the variants in chunks of the given size to limit the memory requirements. Its default value ( <code>NA_integer_</code> ) implies that the whole input VCF file will be read into main memory.                                                                                                                                                                          |
| <code>object</code>            | A <code>VariantFilteringParam</code> object created through <code>VariantFilteringParam()</code> .                                                                                                                                                                                                                                                                                                                                                                                                                                                                           |
| <code>x</code>                 | A <code>VariantFilteringParam</code> object created through <code>VariantFilteringParam()</code> .                                                                                                                                                                                                                                                                                                                                                                                                                                                                           |
| <code>name</code>              | Slot name of a <code>VariantFilteringParam</code> object. Use <code>names()</code> to find out what these slots are.                                                                                                                                                                                                                                                                                                                                                                                                                                                         |

## Details

The class `VariantFilteringParam` serves as a purpose of simplifying the call to the inheritance model function and its subsequent annotation and filtering steps. It also groups all the parameters that the user can customize (i.e newer versions of the annotation packages, when available).

The method `VariantFilteringParam()` creates an `VariantFilteringParam` object used as an input argument to other functions such as `autosomalRecessiveHomozygous()`, etc.

The method `names()` allows one to see the names of the slots from a `VariantFilteringParam` object. Using the `$` operator, one can retrieve the values of these slots in an analogous way to a list.

## Value

An `VariantFilteringParam` object is returned by the method `VariantFilteringParam`.

## Author(s)

D.M. Elurbe, P. Puigdevall and R. Castelo

**Examples**

```

vfpar <- VariantFilteringParam(system.file("extdata", "CEUtrio.vcf.bgz", package="VariantFiltering"),
                             system.file("extdata", "CEUtrio.ped", package="VariantFiltering"),
                             snpdb=character(0), otherAnnotations=character(0))

vfpar
names(vfpar)
vfpar$vcfFiles

```

---

VariantFilteringResults-class

*VariantFilteringResults class*


---

**Description**

The VariantFilteringResults class is used to store the kind of object obtained as a result of an analysis using the functions [unrelatedIndividuals\(\)](#), [autosomalRecessiveHomozygous\(\)](#), [autosomalRecessiveHeterozygous\(\)](#), [autosomalDominant\(\)](#), [deNovo\(\)](#) and [xLinked\(\)](#). Its purpose is to ease the task of filtering and prioritizing the variants annotated by those functions.

**Details**

Variants are stored within a VariantFilteringResults object using a [VRanges](#) object, which also holds the variant annotations in its metadata columns. VariantFiltering adds the following core set of annotations.

**LOCATION** Region where the variant is located (coding, intronic, splice site, promoter, ...) as given by the function [locateVariants\(\)](#) from the VariantAnnotation package.

**LOCSTART** Start position of the variant within the region defined by the LOCATION annotation.

**GENEID** Gene identifier derived with the transcript-centric annotation package given in the txdb argument of the [VariantFilteringParam\(\)](#) function, typically an Entrez Gene identifier.

**GENE** Gene name given by HGNC derived with the gene-centric annotation package given in the orgdb argument of the [VariantFilteringParam\(\)](#) function.

**TYPE** Type of variant, either a single nucleotide variant (SNV), an insertion, a deletion, a multi-nucleotide variant (MNV) or a deletion followed by an insertion (Delins). These types are determined using functions [isSNV\(\)](#), [isInsertion\(\)](#), [isDeletion\(\)](#), [isSubstitution\(\)](#) and [isDelins\(\)](#) from the VariantAnnotation package.

**dbSNP** dbSNP identifier derived by position from the annotation packages given in the snpdb argument of the [VariantFilteringParam\(\)](#) function.

**cDNALOC** Location of the variant along the processed transcript, when the variant belongs to an exonic region.

**CONSEQUENCE** Consequence of the variant when located in the coding region (synonymous, nonsynonymous, missense, nonsense or frameshift) as given by the function [predictCoding\(\)](#) from the VariantAnnotation package.

**TXNAME** Transcript name extracted from the TxDb annotation package given by the txdb argument of the [VariantFilteringParam\(\)](#) function.

**HGVSg** HGVS description of the variant at genomic level.

**HGVSc** HGVS description of the variant at coding level.

**HGVSp** HGVS description of the variant at protein level.

**OMIM** OMIM identifier of the gene associated to the variant derived with the gene-centric annotation package given in the `orgdb` argument of the `VariantFilteringParam()` function.

**AAchangeType** In the case of coding variants, whether the amino acid change is conservative or radical according to the matrix of amino acid biochemical properties given in the argument `radicalAAchangeFilename` of the `VariantFilteringParam()` function.

**SCORE5ssREF** Score for the cryptic 5'ss for the REF allele respect to the ALT allele.

**SCORE5ssALT** Maximum score for a potential cryptic 5'ss created by the ALT allele.

**SCORE5ssPOS** Position of the allele respect to the position of the dinucleotide GT, considering those as positions 1 and 2.

**SCORE3ssREF** Score for the cryptic 3'ss for the REF allele respect to the ALT allele.

**SCORE3ssALT** Maximum score for a potential cryptic 3'ss created by the ALT allele.

**SCORE3ssPOS** Position of the allele respect to the position of the dinucleotide AG, considering those as positions 1 and 2.

## Accessors

A `VariantFilteringResults` has the following set of accessor methods.

`length(x)`: total number of variants stored internally within the `VRanges` object. Note that this number will be typically larger than the number of variants in the input VCF object because each of them is copied for each combination of alternate allele, annotated region and sample.

`param(x)`: returns the `VariantFilteringParam` input parameter object employed in the call that produced the `VariantFilteringResults` object `x`.

`inheritanceModel(x)`: returns the model of inheritance employed in the call that produced the `VariantFilteringResults` object `x`.

`samples(object)`: active samples from which the current filtered variants were derived. If the `x` was obtained with `unrelatedIndividuals()`, then the `replace` method `samples(object)<-` can be used to restrict the subset of active samples. In every other case (`autosomalDominant()`, etc. ) active samples cannot be changed.

`resetSamples(object)`: set back as active samples the initial set of samples specified in the input parameter object.

`sog(x)`: Sequence Ontology (SO) graph (actually, an acyclic digraph) returned as a `graphNEL` object, whose vertices are SO terms, edges represent ontology relationships and vertex attributes `vcfIdx` and `varIdx` contain what variants are annotated to each SO term. These annotations can be directly retrieved from the SO graph with the `nodeData()` function from the `graph` package. The `summary()` function described in this manual page allows one to tally the number of variants in each SO term throughout the entire SO hierarchy.

`bamFiles(x)`: access and update the `BamViews` object containing references to BAM files from which the input VCF files were derived. Initially this is empty.

`allVariants(x, groupBy="sample")`: returns a `VRangesList` object with all variants grouped by default by sample. Using the argument `groupBy` we can specify any metadata column to be used to group variants. If the value given to `groupBy` does not correspond to any such columns, a `VRanges` object with all variants together is returned.

`filteredVariants(x, groupBy="sample")`: it works like `allVariants(x)` but instead of returning all variants, it returns only those who pass the active filters; see `filters()` and `cutoffs()` below.

## Filters and cutoffs

The variants contained in a `VariantFilteringResults` object can be filtered using the `FilterRules` mechanism, defined in the `S4Vectors` package, by using the functions `filters()` and `cutoffs()` described below. There are additional functions, also described in this section, to facilitate this task on the set of core annotations provided by `VariantFiltering`.

`filters(x)`: get the current `FilterRules` object that defines the available set of filter criteria that one can use to filter the variants contained in `x`. This can also be used as a replacement function `filters(x)<-` to update this set of filters. The actual filtering is done when calling the function `filteredVariants()`.

`filtersMetadata(x)`: metadata about the available filters.

`cutoffs(x)`: get cutoffs from the available filters.

`change(x, cutoff)<-`: change cutoffs from the available filters. Here, argument `x` is a `CutoffsList` object given by the method `cutoffs()`, and argument `cutoff` is a character string with the name of the cutoff.

`softFilterMatrix(x)`: get and update the variant by filter matrix; see `softFilterMatrix()` in the `VariantAnnotation` package.

`dbSNPpresent(x)`: flag whether to filter variants present or absent from dbSNP (NA -do not filter-, "Yes", "No").

`variantType(x)`: filter by type of variant ("SNV", "Insertion", "Deletion", "MNV", "Delins").

`variantLocation(x)`: filter by variant location ("coding", "intron", "threeUTR", "fiveUTR", "intergenic", "spliceSite", "promoter").

`variantConsequence(x)`: filter by variant consequence ("synonymous", "nonsynonymous", "frameshift", "nonsense", "not translated").

`aaChangeType(x)`: filter by type of change of amino acid ("Radical", "Conservative").

`OMIMpresent(x)`: flag whether to filter variants whose associated genes are present or absent from OMIM (NA -do not filter-, "Yes", "No").

`naMAF(x)`: flag whether NA maximum MAF values should be included in the filtered variants.

`maxMAF(x)`: maximum MAF value that a variant may meet among the selected populations.

`minPhastCons(x)`: minimum phastCons score for nucleotide conservation (NA -do not filter-, [0-1]).

`minPhylostratum(x)`: minimum phylostratum for gene conservation (NA -do not filter-, [1-20]).

`MAFpop(x)`: selection of populations to use when filtering by maximum MAF value.

`minScore5ss(x)`: minimum weight matrix score on a cryptic 5'ss. NA indicates this filter is not applied.

`minScore3ss(x)`: minimum weight matrix score on a cryptic 3'ss. NA indicates this filter is not applied.

`minCUFC(x)`: minimum absolute codon-usage log2 fold-change.

## Summarization, visualization and reporting

The following functions help in summarizing, visualizing and reporting the filtered variants.

`summary(object, method=c("SO", "SOfull", "bioc"))`: tally the current filtered set of variants to features. By default, features are Sequence Ontology (SO) terms to which variants are annotated by `VariantFiltering`. The method argument allows the user to change this default setting to tallying throughout the entire SO hierarchy. Both options, `SO` and `SOfull` can be used in combination with the cutoff `SOterms`; see the vignette. The option `method="bioc"` considers as features the regions and consequences annotated by functions `locateVariants()` and `predictCoding()` from the `VariantAnnotation` package. The result is returned as a `data.frame` object.

`plot(x, what, sampleName, flankingNt=20, showAlnNtCutoff=200, isPaired=FALSE, ...)`: Plot variants using the `Gviz` package. The argument `what` can be either a character vector specifying gene or variant identifiers or a chromosome name, or a `GRanges` object specifying a genomic region. The argument `sampleName` is optional and allows the user to plot the aligned reads and coverage from a specific sample, located in the plotted region, when the corresponding BAM file has been linked to the object with `bamFiles()`. The argument `flankingNt` is a number of nucleotides to extend the plotting region derived from the argument `what`. The argument `showAlnNtCutoff` is the region size cutoff below which it will be attempted to plot the aligned reads. The argument `isPaired` is passed directly to the `Gviz` function `AlignmentsTrack()` which streams over the BAM file to plot the reads and sets whether the BAM file contains single (default) or paired-end reads. Further arguments in `...` are passed to the `Gviz` function `plotTracks()` and can be used to fine-tune the final plot; see the vignette of `Gviz` to find out what these arguments are.

`reportVariants(x, type=c("shiny", "csv", "tsv"), file=NULL)`: Builds a report from the `VariantFilteringResult` object `x`. Using the type argument, the report can take the form of a flat file in CSV or TSV format or a web shiny app (default) that enables applying functional annotation filters in an interactive manner.

When the shiny app is closed this method returns a `VariantFilteringResult` object with the corresponding filters switched on or off according to how the app has been interactively used.

## Author(s)

R. Castelo

## Examples

```
## Not run:
library(VariantFiltering)

CEUvcf <- file.path(system.file("extdata", package="VariantFiltering"),
                    "CEUtrio.vcf.gz")
CEUped <- file.path(system.file("extdata", package="VariantFiltering"),
```

```

                                "CEUtrio.ped")
param <- VariantFilteringParam(vcfFileNames=CEUvcf, pedFileName=CEUped)
reHo <- autosomalRecessiveHomozygous(param)
naMAF(reHo) <- FALSE
maxMAF(reHo) <- 0.05
reHo
head(filteredVariants(reHo))
reportVariants(reHo, type="csv", file="reHo.csv")

## End(Not run)

```

---

|                   |                                                                                     |
|-------------------|-------------------------------------------------------------------------------------|
| VariantType-class | <i>FiveSpliceSiteVariants and ThreeSpliceSiteVariants subclasses of VariantType</i> |
|-------------------|-------------------------------------------------------------------------------------|

---

## Description

VariantType subclasses specify the type of variant to be located with the function `locateVariants()` from the VariantAnnotation package. Here we describe two such subclasses defined within the VariantFiltering package: `FiveSpliceSiteVariants` and `ThreeSpliceSiteVariants`. For further details on the [VariantType-class](#) please consult the corresponding manual page in the VariantAnnotation package.

## Usage

```

FiveSpliceSiteVariants(minIntronLength=20L, upstream = 3L, downstream = 4L)
ThreeSpliceSiteVariants(minIntronLength=20L, upstream = 18L, downstream = 3L)

```

## Arguments

`minIntronLength`

Minimum intron length in nucleotides, by default `minIntronLength=20L`. Splice sites annotated within introns smaller than this length will not be used to annotate variants.

`upstream, downstream`

Single integer values representing the number of nucleotides upstream and downstream of the 5' and 3' dinucleotide ends of introns. By default, the constructor function `FiveSpliceSiteVariants` has values `upstream=3L` and `downstream=4L`, corresponding to 3 nucleotides upstream and 4 nucleotides downstream of the first dinucleotide of the intron, while `ThreeSpliceSiteVariants` has values `upstream=18L` and `downstream=3L`, corresponding to 18 nucleotides upstream and 4 nucleotides downstream of the last dinucleotide of the intron.

## Details

The subclasses `FiveSpliceSiteVariants` and `ThreeSpliceSiteVariants` can be used as in the `region` argument to the `locateVariants()` function from the VariantAnnotation package or as values in the `regionAnnotations` argument to the `VariantFilteringParam()` function that

constructs a [VariantFilteringParam-class](#) object. These two subclasses allow the package VariantFiltering and the function [locateVariants\(\)](#) in the VariantAnnotation package, to annotate variants located within a splice site region defined by the user. The boundaries of this region as well as the minimum intron length can be defined with the following arguments to the constructor functions:

### Accessors

In the following code, x is a PromoterVariants or a AllVariants object.

`minIntronLength(x), minIntronLength(x) <- value:` Gets or sets the minimum number of nucleotides long of introns to be considered in the annotation of splice sites.

`upstream(x), upstream(x) <- value:` Gets or sets the number of nucleotides upstream of the 5' and 3' end of the intron, excluding the first and last dinucleotide, respectively.

`downstream(x), downstream(x) <- value:` Gets or sets the number of nucleotides downstream of the 5' and 3' end of the intron, excluding the first and last dinucleotide, respectively.

### Author(s)

Robert Castelo <[robert.castelo@upf.edu](mailto:robert.castelo@upf.edu)>

### See Also

- The [VariantType-class](#) man page in the VariantAnnotation package.

### Examples

```
FiveSpliceSiteVariants()
ThreeSpliceSiteVariants()
```

---

|                    |                            |
|--------------------|----------------------------|
| WeightMatrix-class | <i>Weight matrix class</i> |
|--------------------|----------------------------|

---

### Description

Class for storing weight matrices that VariantFiltering uses to score potential cryptic splice sites.

### Usage

```
## S4 method for signature 'WeightMatrix'
width(x)
## S4 method for signature 'WeightMatrix'
conservedPositions(x)
## S4 method for signature 'WeightMatrix'
wmName(x)
## S4 method for signature 'WeightMatrix'
wmFilename(x)
## S4 method for signature 'WeightMatrix'
```

```

wmLocations(x)
## S4 method for signature 'WeightMatrix'
wmStrictLocations(x)
## S4 method for signature 'WeightMatrix'
show(object)
## S4 method for signature 'WeightMatrix,DNAStringSet'
wmScore(object, dnaseqs)
## S4 method for signature 'WeightMatrix,character'
wmScore(object, dnaseqs)
## S4 method for signature 'character,DNAStringSet'
wmScore(object, dnaseqs, locations, strictLocations)
## S4 method for signature 'character,character'
wmScore(object, dnaseqs, locations, strictLocations)

```

### Arguments

|                              |                                                                                                                                                                                                               |
|------------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <code>x</code>               | A <code>WeightMatrix</code> object.                                                                                                                                                                           |
| <code>object</code>          | A <code>WeightMatrix</code> object or the file name of a weight matrix.                                                                                                                                       |
| <code>dnaseqs</code>         | Either a vector of character strings a <code>DNAStringSet</code> object, both of which store nucleotide sequences to be scored using the input <code>WeightMatrix</code> object.                              |
| <code>locations</code>       | Character vector of the annotated locations to variants under which the weight matrix will be used for scoring binding sites. The possible values can be obtained by typing <code>variantLocations()</code> . |
| <code>strictLocations</code> | Logical vector flagging whether the weight matrix should be scoring binding sites strictly within the boundaries of the given locations.                                                                      |

### Details

The `WeightMatrix` class and associated methods serve the purpose of enabling the `VariantFiltering` package to score synonymous and intronic genetic variants for potential cryptic splice sites. The class and the methods, however, are exposed to the end user since they could be useful for other analysis purposes.

The `VariantFiltering` package contains two weight matrices, one for 5'ss and another for 3'ss, which have been built by a statistical method that accounts for dependencies between the splice site positions, minimizing the rate of false positive predictions. The method concretely builds these models by inclusion-driven learning of Bayesian networks and further details can be found in the paper of Castelo and Guigo (2004).

The function `readWm()` reads a weight matrix stored in a text file in a particular format and returns a `WeightMatrix` object. See the `.ibn` files located in the `extdata` folder of the `VariantFiltering` package, as an example of this format that is specifically designed to enable the storage of weights that may depend on the occurrence of nucleotides at other positions on the matrix.

Next to this specific weight matrix format, the function `readWm()` can also read the MEME motif format specified at <http://meme-suite.org/doc/meme-format.html>. Under this format, this function reads only currently one matrix per file and when values correspond to probabilities (specified under the motif letter-probability matrix line) they are automatically converted to weights

by either using the background frequencies specified in the background letter frequencies line or using an uniform distribution otherwise.

The method `wmScore()` scores one or more sequences of nucleotides using the input `WeightMatrix` object. When the input object is the file name of a weight matrix, the function `readWm()` is called to read first that weight matrix and internally create a `WeightMatrix` object. This way to call `wmScore()` is required when using it in parallel since currently `WeightMatrix` objects are not serializable.

If the sequences are longer than the width of the weight matrix, this function will score every possible site within those sequences. It returns a list where each element is a vector with the calculated scores of the corresponding DNA sequence. When the scores cannot be calculated because of a conserved position that does not occur in the sequence (i.e., absence of a GT dinucleotide with the 5'ss weight matrix), it returns NA as corresponding score value.

The method `width()` takes a `WeightMatrix` object as input and returns the number of positions of the weight matrix.

The method `conservedPositions()` takes a `WeightMatrix` object as input and returns the number of fully conserved positions in the weight matrix.

## Value

.

## Author(s)

R. Castelo

## References

Castelo, R and Guigo, R. Splice site identification by idIBNs. *Bioinformatics*, 20(1):i69-i76, 2004.

## Examples

```
wm <- readWm(file.path(system.file("extdata", package="VariantFiltering"), "hsap.donors.hcmc10_15_1.ibn"),
             locations="fiveSpliceSite", strictLocations=TRUE)

wm
wmFilename(wm)
width(wm)
wmName(wm)
wmLocations(wm)
wmStrictLocations(wm)
conservedPositions(wm)
wmScore(wm, "CAGGTAGGA")
wmScore(wm, "CAGGAAGGA")
wmScore(wm, "CAGGTCCTG")
wmScore(wm, "CAGGTCGTGGAG")
```

xLinked

*X-Linked inheritance analysis***Description**

This function identifies variants that appear only in the X chromosome of the unaffected females as heterozygous, don't appear in the unaffected males analyzed and finally are present (as homozygous) in the affected male(s).

**Usage**

```
## S4 method for signature 'VariantFilteringParam'
xLinked(param,
        svparam=ScanVcfParam(),
        use=c("everything", "complete.obs", "all.obs"),
        BPPARAM=bpparam("SerialParam"))
```

**Arguments**

|         |                                                                                                                                                                                                                                                                                                                                                                                                                   |
|---------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| param   | A <a href="#">VariantFilteringParam</a> object built from a multisample VCF file with at least one affected individual and zero or more unaffected ones, and from a PED file specifying the family relationships among individuals as well as their gender and phenotype status (affected or unaffected).                                                                                                         |
| svparam | An instance of a <a href="#">ScanVcfParam</a> object to enable analyzing a subset of variants and samples. This object is passed internally to a call to the <a href="#">readVcf()</a> function in the <a href="#">VariantAnnotation</a> package, see its help page for a complete description of this functionality.                                                                                             |
| use     | character string specifying the policy to apply on missing genotypes when comparing them. This policy can be either "everything" (default), "complete.obs" or "all.obs". The default policy ("everything") will propagate NA truth values using the behavior of the R logical operators, with the exception that when the final truth value associated with a variant is NA, the variant is ultimately discarded. |
| BPPARAM | An object of class <a href="#">BiocParallelParam</a> specifying parameters related to the parallel execution of some of the tasks and calculations within this function. See function <a href="#">bpparam()</a> from the <a href="#">BiocParallel</a> package.                                                                                                                                                    |

**Details**

This function requires as an input a [VariantFilteringParam](#) class object built from an input multisample VCF file, along with a PED file.

**Value**

An object of class [VariantFilteringResults](#) including functional annotations on all selected variants.

**Author(s)**

Dei M. Elurbe and R. Castelo

**References**

Elurbe D.M., Mila, M., Castelo, R. The VariantFiltering package, in preparation.

**See Also**

[autosomalRecessiveHomozygous](#) [autosomalRecessiveHeterozygous](#) [autosomalDominant](#) [deNovo](#)  
[unrelatedIndividuals](#) [VariantFilteringResults](#)

**Examples**

```
## Not run:

## This actually won't run b/c in this trio de descendant is a female
CEUvcf <- file.path(system.file("extdata", package="VariantFiltering"), "CEUtrio.vcf.bgz")
CEUped <- file.path(system.file("extdata", package="VariantFiltering"), "CEUtrio.ped")
param <- VariantFilteringParam(vcfFilename=CEUvcf,
                              pedFilename=CEUped)
xlid <- xLinked(param)
xlid

## End(Not run)
```

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