

Package ‘gwascat’

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Title representing and modeling data in the EMBL-EBI GWAS catalog

Version 2.36.0

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Description Represent and model data in the EMBL-EBI GWAS catalog.

Enhances SNPlocs.Hsapiens.dbSNP144.GRCh37

Depends R (>= 3.5.0), methods

Imports S4Vectors (>= 0.9.25), IRanges, GenomeInfoDb, GenomicRanges (>= 1.29.6), GenomicFeatures, readr, Biostrings, AnnotationDbi, BiocFileCache, snpStats, VariantAnnotation, AnnotationHub

Suggests DO.db, DT, knitr, RBGL, testthat, rmarkdown, dplyr, Gviz, Rsamtools, rtracklayer, graph, ggbio, DelayedArray, TxDb.Hsapiens.UCSC.hg19.knownGene, org.Hs.eg.db, BiocStyle

VignetteBuilder knitr

Maintainer VJ Carey <stvjc@channing.harvard.edu>

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as_GRanges

produce a GRanges from gwascat tibble

Description

produce a GRanges from gwascat tibble

Usage

```
as_GRanges(
  x,
  short = TRUE,
  for_short = c("PUBMEDID", "DATE", "DISEASE/TRAIT", "SNPS"),
  genome_tag = "GRCh38"
)
```

Arguments

| | |
|------------|--|
| x | a tibble from 'get_cached_gwascat()' |
| short | logical(1) if TRUE only keep selected columns in mcols |
| for_short | character() column names to keep in mcols |
| genome_tag | character(1) defaults to "GRCh38" |

 bindcadd_snv

bind CADD scores of Kircher et al. 2014 to a GRanges instance

Description

bind CADD scores of Kircher et al. 2014 to a GRanges instance; by default will use HTTP access at UW

Usage

```
bindcadd_snv(
  gr,
  fn = "http://krishna.gs.washington.edu/download/CADD/v1.0/1000G.tsv.gz"
)
```

Arguments

| | |
|----|--|
| gr | query ranges to which CADD scores should be bound |
| fn | path to Tabix-indexed bgzipped TSV of CADD as distributed at krishna.gs.washington.edu on 1 April 2014 |

Details

joins CADD fields at addresses that match query; the CADD fields for query ranges that are not matched are set to NA

Value

GRanges instance with additional fields as obtained in the CADD resource

Note

This software developed in part with support from Genentech, Inc.

Author(s)

VJ Carey <stvjc@channing.harvard.edu>

References

M Kircher, DM Witten, P Jain, BJ O’Roak, GM Cooper, J Shendure, A general framework for estimating the relative pathogenicity of human genetic variants, Nature Genetics Feb 2014, PMID 24487276

Examples

```
## Not run:
data(ebicat_2020_04_30)
g2 = as(ebicat_2020_04_30, "GRanges")
# would need to lift over here
bindcadd_snv( g2[which(seqnames(g2)=="chr2")][1:20] )

## End(Not run)
```

| | |
|---------|---|
| chklocs | <i>return TRUE if all named SNPs with locations in both the SNPlocs package and the gwascat agree</i> |
|---------|---|

Description

return TRUE if all named SNPs with locations in both the SNPlocs package and the gwascat agree

Usage

```
chklocs(chrtag = "20", gwwl = gwrngs19)
```

Arguments

| | |
|--------|----------------------------------|
| chrtag | character, chromosome identifier |
| gwwl | instance of {gwaswloc} |

| | |
|-------------------|---|
| ebicat_2020_04_30 | <i>serialized gwaswloc instance from april 30 2020, sample of 50000 records</i> |
|-------------------|---|

Description

serialized gwaswloc instance from april 30 2020, sample of 50000 records

Usage

ebicat_2020_04_30

Format

gwaswloc instance

| | |
|-------|--------------------------------------|
| g17SM | <i>SnpMatrix instance from chr17</i> |
|-------|--------------------------------------|

Description

SnpMatrix instance from chr17

Usage

g17SM

Format

snpStats SnpMatrix instance

| | |
|----------|-----------------------------------|
| getRsids | <i>generic snp name retrieval</i> |
|----------|-----------------------------------|

Description

generic snp name retrieval

Usage

getRsids(x)

Arguments

x gwaswloc

getRsids,gwaswloc-method
specific snp name retrieval

Description

specific snp name retrieval

Usage

```
## S4 method for signature 'gwaswloc'  
getRsids(x)
```

Arguments

x gwaswloc

getTraits *generic trait retrieval*

Description

generic trait retrieval

Usage

```
getTraits(x)
```

Arguments

x gwaswloc

```
getTraits,gwaswloc-method
      specific trait retrieval
```

Description

specific trait retrieval

Usage

```
## S4 method for signature 'gwaswloc'
getTraits(x)
```

Arguments

```
x          gwaswloc
```

```
get_cached_gwascat    use BiocFileCache to retrieve and keep an image of the tsv file distributed by EBI
```

Description

use BiocFileCache to retrieve and keep an image of the tsv file distributed by EBI

Usage

```
get_cached_gwascat(
  url = "http://www.ebi.ac.uk/gwas/api/search/downloads/alternative",
  cache = BiocFileCache::BiocFileCache(),
  refresh = FALSE,
  ...
)
```

Arguments

```
url          character(1) url to use
cache        BiocFileCache::BiocFileCache instance
refresh      logical(1) force download and recaching
...          passed to bfcadd
```

Value

a tibble as produced by readr::read_tsv, with attributes extractDate (as recorded in cache as 'access_time', and problems (a tibble returned by read_tsv).

Note

will If query of cache with 'ebi.ac.uk/gwas' returns 0-row tibble, will populate cache with bfcadd. Uses readr::read_tsv on cache content to return tibble. The etag field does not seem to be used at EBI, thus user must check for updates.

`gg17N`*genotype matrix from chr17 1000 genomes*

Description

genotype matrix from chr17 1000 genomes

Usage`gg17N`**Format**

matrix

Examples

```
data(gg17N)
gg17N[1:4, 1:4]
```

`gr6.0_hg38`*image of locon6 in GRanges, lifted over to hg38*

Description

image of locon6 in GRanges, lifted over to hg38

Usage`gr6.0_hg38`**Format**

GRanges instance

| | |
|-----------|--|
| gw6.rs_17 | <i>character vector of rs numbers for SNP on chr17</i> |
|-----------|--|

Description

character vector of rs numbers for SNP on chr17

Usage

```
gw6.rs_17
```

Format

character vector

| | |
|-------------------|---|
| gwascat_from_AHub | <i>grab an image of EBI GWAS catalog from AnnotationHub</i> |
|-------------------|---|

Description

grab an image of EBI GWAS catalog from AnnotationHub

Usage

```
gwascat_from_AHub(tag = "AH91571", simple = FALSE, fixNonASCII = TRUE)
```

Arguments

| | |
|-------------|--|
| tag | character(1) defaults to "AH91571" which is the 3.30.2021 image |
| simple | logical(1) if TRUE, just returns data.frame as retrieved from EBI; defaults to FALSE |
| fixNonASCII | logical(1) if TRUE, use iconv to identify and eliminate non-ASCII content |

Value

If 'simple', a data.frame is returned based on TSV data produced by EBI. Otherwise, non-ASCII content is processed according to the value of 'fixNonASCII' and a 'gwaswloc' instance is returned, which has a concise show method. This can be coerced to a simple GRanges instance with as(..., "GRanges"). The reference build is GRCh38.

Examples

```
gwc = gwascat_from_AHub()
gwc
```

| | |
|------------|--|
| gwastagger | <i>GRanges with LD information on 9998 SNP</i> |
|------------|--|

Description

GRanges with LD information on 9998 SNP

Usage

gwastagger

Format

GRanges

| | |
|----------------|--|
| gwaswloc-class | <i>container for gwas hit data and GRanges for addresses</i> |
|----------------|--|

Description

container for gwas hit data and GRanges for addresses

| | |
|---------------|--|
| gwcatsnapshot | <i>use AnnotationHub snapshot as basis for gwaswloc structure creation</i> |
|---------------|--|

Description

use AnnotationHub snapshot as basis for gwaswloc structure creation

Usage

gwcatsnapshot(x, fixNonASCII = TRUE)

Arguments

| | |
|-------------|---|
| x | inherits from data.frame, with columns consistent with EBI table |
| fixNonASCII | logical(1) if TRUE, use iconv to replace non-ASCII character, important for CMD check but perhaps not important for applied use |

Examples

```

ah = AnnotationHub::AnnotationHub()
entitytab = AnnotationHub::query(ah, "gwascatData")
cand = names(entitytab)[1]
stopifnot(nchar(cand)>0)
tab = ah[[cand]]
gww = gwascat_snapshot(tab)
gww
length(gww)

```

gwcecx2gviz

Prepare salient components of GWAS catalog for rendering with Gviz

Description

Prepare salient components of GWAS catalog for rendering with Gviz

Usage

```

gwcecx2gviz(
  basegr,
  contextGR = GRanges(seqnames = "chr17", IRanges::IRanges(start = 37500000, width =
    1e+06)),
  txrefobj = TxDb.Hsapiens.UCSC.hg19.knownGene::TxDb.Hsapiens.UCSC.hg19.knownGene,
  genome = "hg19",
  genesymobj = org.Hs.eg.db::org.Hs.eg.db,
  plot.it = TRUE,
  maxmlp = 25
)

```

Arguments

| | |
|------------|--|
| basegr | gwaswloc instance containing information about GWAS in catalog |
| contextGR | A GRanges instance delimiting the visualization in genomic coordinates |
| txrefobj | a TxDb instance |
| genome | character tag like 'hg19' |
| genesymobj | an OrgDb instance |
| plot.it | logical, if FALSE, just return list |
| maxmlp | maximum value of $-10 \log p$ – winsorization of all larger values is performed, modifying the contents of Pvalue_mlogp in the elementMetadata for the call |

Examples

```

data(ebicat_2020_04_30)
# GenomeInfoDb::seqlevelsStyle(ebicat_2020_04_30) = "UCSC" # no more
GenomeInfoDb::seqlevels(ebicat_2020_04_30) = paste0("chr", GenomeInfoDb::seqlevels(ebicat_2020_04_30))
gwcecx2gviz(ebicat_2020_04_30)

```

ldtagr *expand a list of variants by including those in a VCF with LD exceeding some threshold; uses snpStats ld()*

Description

expand a list of variants by including those in a VCF with LD exceeding some threshold; uses snpStats ld()

Usage

```
ldtagr(
  snprng,
  tf,
  samples,
  genome = "hg19",
  lbmaf = 0.05,
  lbR2 = 0.8,
  radius = 1e+05
)
```

Arguments

| | |
|---------|---|
| snprng | a named GRanges for a single SNP. The name must correspond to the name that will be assigned by genotypeToSnpMatrix (from VariantTools) to the corresponding column of a SnpMatrix. |
| tf | TabixFile instance pointing to a bgzipped tabix-indexed VCF file |
| samples | a vector of sample identifiers, if excluded, all samples used |
| genome | tag like 'hg19' |
| lbmaf | lower bound on variant MAF to allow consideration |
| lbR2 | lower bound on R squared for regarding SNP to be incorporated |
| radius | radius of search in bp around the input range |

Value

a GRanges with names corresponding to 'new' variants and mcols fields 'paramRangeID' (base variant input) and 'R2'

Note

slow but safe approach. probably a matrix method could be substituted using the nice sparse approach already in snpStats

Author(s)

VJ Carey

Examples

```

cand = GenomicRanges::GRanges("1", IRanges::IRanges(113038694, width=1))
names(cand) = "rs883593"
requireNamespace("VariantAnnotation")
expath = dir(system.file("vcf", package="gwascat"), patt="*exon.*gz$", full=TRUE)
tf = Rsamtools::TabixFile(expath)
ldtagr( cand, tf, lbR2 = .8)

```

| | |
|--------|------------------------------------|
| locon6 | <i>location data for 10000 SNP</i> |
|--------|------------------------------------|

Description

location data for 10000 SNP

Usage

locon6

Format

data.frame, coordinates are hg19

| | |
|------------|---|
| locs4trait | <i>get locations for SNP affecting a selected trait</i> |
|------------|---|

Description

get locations for SNP affecting a selected trait

Usage

```
locs4trait(gwvl, trait, tag = "DISEASE/TRAIT")
```

Arguments

| | |
|-------|---|
| gwvl | instance of {gwaswloc} |
| trait | character, name of trait |
| tag | character, name of field to be used for trait enumeration |

| | |
|-------|--------------------------------------|
| low17 | <i>SnpMatrix instance from chr17</i> |
|-------|--------------------------------------|

Description

SnpMatrix instance from chr17

Usage

low17

Format

snpStats SnpMatrix instance

| | |
|--------------------|---|
| makeCurrentGwascat | <i>read NHGRI GWAS catalog table and construct associated GRanges instance records for which clear genomic position cannot be determined are dropped from the ranges instance an effort is made to use reasonable data types for GRanges metadata, so some qualifying characters such as (EA) in Risk allele frequency field will simply be omitted during coercion of contents of that field to numeric.</i> |
|--------------------|---|

Description

read NHGRI GWAS catalog table and construct associated GRanges instance records for which clear genomic position cannot be determined are dropped from the ranges instance an effort is made to use reasonable data types for GRanges metadata, so some qualifying characters such as (EA) in Risk allele frequency field will simply be omitted during coercion of contents of that field to numeric.

Usage

```
makeCurrentGwascat(
  table.url = "http://www.ebi.ac.uk/gwas/api/search/downloads/alternative",
  fixNonASCII = TRUE,
  genome = "GRCh38",
  withOnt = TRUE
)
```

Arguments

| | |
|--------------------------|---|
| <code>table.url</code> | string identifying the .txt file curated at EBI/EMBL |
| <code>fixNonASCII</code> | logical, if TRUE, non-ASCII characters as identified by <code>iconv</code> will be replaced by asterisk |
| <code>genome</code> | character string: 'GRCh38' is default and yields current image as provided by EMBL/EBI; 'GRCh37' yields a realtime liftOver to hg19 coordinates, via AnnotationHub storage of the chain files. Any other value yields an error. |
| <code>withOnt</code> | logical indicating whether 'alternative' (ontology-present, includes repetition of loci with one:many ontological mapping) or 'full' (ontology-absent, one record per locus report) version of distributed table |

Value

a slightly extended `GRanges` instance, with class name 'gwaswloc'; the purpose of the introduction of this class is to support a concise `show` method that does not produce very long lines owing to large numbers of fields in the `mcols` component.

Note

'`readr::read_tsv`' records problems when some records have field contents that are inconsistent with the column specification. This information can be retrieved from the metadata slot of the returned object, as noted in a message produced when this function is run.

Author(s)

VJ Carey

Examples

```
# if you have good internet access
if (interactive()) {
  newcatr = makeCurrentGwascat()
  newcatr
}
```

| | |
|--------------|---|
| obo2graphNEL | <i>convert a typical OBO text file to a graphNEL instance (using Term elements)</i> |
|--------------|---|

Description

convert a typical OBO text file to a graphNEL instance (using Term elements)

Usage

```

obo2graphNEL(
  obo = "human-phenotype-ontology.obo",
  kill = "\\[Typedef\\]",
  killTrailSp = TRUE
)

```

Arguments

| | |
|-------------|---|
| obo | string naming a file in OBO format |
| kill | entity types to be excluded from processing – probably this should be in a 'keep' form, but for now this works. |
| killTrailSp | In the textual version of EFO ca. Aug 2015, there is a trailing blank in the tag field defining EFO:0000001, which is not present in references to this term. Set this to TRUE to eliminate this, or graphNEL construction will fail to validate. |

Details

Very rudimentary list and grep operations are used to retain sufficient information to map the DAG to a graphNEL, using formal term identifiers as node names and 'is-a' relationships as edges, and term names and other metadata are assigned to nodeData components.

Value

a graphNEL instance

Note

The OBO for Human Disease ontology is serialized as text with this package.

Author(s)

VJ Carey <stvjc@channing.harvard.edu>

References

For use with human disease ontology, http://www.obofoundry.org/cgi-bin/detail.cgi?id=disease_ontology

Examples

```

data(efo.obo.g)
requireNamespace("graph")
hn = graph::nodes(efo.obo.g)[1:5]
hn
graph::nodeData(efo.obo.g, hn[5])

```

process_gwas_dataframe

convert GWAS catalog data.frame to gwaswloc, a GRanges extension with simple show method

Description

convert GWAS catalog data.frame to gwaswloc, a GRanges extension with simple show method

Usage

```
process_gwas_dataframe(df)
```

Arguments

df data.frame

riskyAlleleCount *given a matrix of subjects x SNP calls, count number of risky alleles*

Description

given a matrix of subjects x SNP calls, count number of risky alleles for various conditions, relative to NHGRI GWAS catalog

Usage

```
riskyAlleleCount(
  callmat,
  matIsAB = TRUE,
  chr,
  gwwl,
  snpap = "SNPlocs.Hsapiens.dbSNP144.GRCh37",
  gencode = c("A/A", "A/B", "B/B")
)
```

Arguments

| | |
|---------|--|
| callmat | matrix with subjects as rows, SNPs as columns; entries can be generic A/A, A/B, B/B, or specific nucleotide calls |
| matIsAB | logical, FALSE if nucleotide codes are present, TRUE if generic call codes are present; in the latter case, gwascat::ABmat2nuc will be run |
| chr | code for chromosome, should work with the SNP annotation getSNPlocs function, so likely "ch[nn]" |
| gwwl | an instance of {gwaswloc} |
| snpap | name of a Bioconductor SNPlocs.Hsapiens.dbSNP.* package |
| gencode | codes used for generic SNP call |

Value

matrix with rows corresponding to subjects , columns corresponding to SNP

Examples

```
## Not run:
data(gg17N) # translated from GGdata chr 17 calls using ABmat2nuc
data(ebicat37)
library(GenomeInfoDb)
seqlevelsStyle(ebicat37) = "UCSC"
h17 = riskyAlleleCount(gg17N, matIsAB=FALSE, chr="ch17", gwwl=ebicat37)
h17[1:5,1:5]
table(as.numeric(h17))

## End(Not run)
```

 si.hs.37

Seqinfo for GRCh37

Description

Seqinfo for GRCh37

Usage

si.hs.37

Format

GenomeInfoDb Seqinfo instance

 si.hs.38

Seqinfo for GRCh38

Description

Seqinfo for GRCh38

Usage

si.hs.38

Format

GenomeInfoDb Seqinfo instance

subsetByChromosome *generic trait subsetting*

Description

generic trait subsetting

Usage

```
subsetByChromosome(x, ch)
```

Arguments

| | |
|----|---------------------------------|
| x | gwaswloc |
| ch | character vector of chromosomes |

subsetByChromosome, gwaswloc-method
specific trait subsetting

Description

specific trait subsetting

Usage

```
## S4 method for signature 'gwaswloc'  
subsetByChromosome(x, ch)
```

Arguments

| | |
|----|---------------------------------|
| x | gwaswloc |
| ch | character vector of chromosomes |

| | |
|----------------|---------------------------------|
| subsetByTraits | <i>generic trait subsetting</i> |
|----------------|---------------------------------|

Description

generic trait subsetting

Usage

```
subsetByTraits(x, tr)
```

Arguments

| | |
|----|----------------------------|
| x | gwaswloc |
| tr | character vector of traits |

| | |
|--------------------------------|----------------------------------|
| subsetByTraits,gwaswloc-method | <i>specific trait subsetting</i> |
|--------------------------------|----------------------------------|

Description

specific trait subsetting

Usage

```
## S4 method for signature 'gwaswloc'  
subsetByTraits(x, tr)
```

Arguments

| | |
|----|----------------------------|
| x | gwaswloc |
| tr | character vector of traits |

| | |
|-----------|-----------------------------------|
| topTraits | <i>operations on GWAS catalog</i> |
|-----------|-----------------------------------|

Description

operations on GWAS catalog

Usage

```
topTraits(gwwl, n = 10, tag = "DISEASE/TRAIT")
```

Arguments

| | |
|------|---|
| gwwl | instance of {gwaswloc} |
| n | numeric, number of traits to report |
| tag | character, name of field to be used for trait enumeration |

Value

topTraits returns a character vector of most frequently occurring traits in the database

locs4trait returns a {gwaswloc} object with records defining associations to the specified trait

chklocs returns a logical that is TRUE when the asserted locations of SNP in the GWAS catalog agree with the locations given in the dbSNP package SNPlocs.Hsapiens.dbSNP144.GRCh37

Author(s)

VJ Carey <stvjc@channing.harvard.edu>

Examples

```
data(ebicat_2020_04_30)
topTraits(ebicat_2020_04_30)
```

| | |
|------------|--|
| traitsManh | <i>use ggbio facilities to display GWAS results for selected traits in genomic coordinates</i> |
|------------|--|

Description

use ggbio facilities to display GWAS results for selected traits in genomic coordinates

Usage

```
traitsManh(
  gwr,
  selr = GRanges(seqnames = "chr17", IRanges(3e+07, 5e+07)),
  traits = c("Asthma", "Parkinson's disease", "Height", "Crohn's disease"),
  truncmlp = 25,
  ...
)
```

Arguments

| | |
|-----------------------|--|
| <code>gwr</code> | GRanges instance as managed by the gwaswloc container design, with Disease.Trait and Pvalue_mlog among elementMetadata columns |
| <code>selr</code> | A GRanges instance to restrict the gwr for visualization. Not tested for noncontiguous regions. |
| <code>traits</code> | Character vector of traits to be exhibited; GWAS results with traits not among these will be labeled "other". |
| <code>truncmlp</code> | Maximum value of $-\log_{10} p$ to be displayed; in the raw data this ranges to the hundreds and can cause bad compression. |
| <code>...</code> | not currently used |

Details

uses a ggbio autoplot

Value

autoplot value

Note

An xlab is added, concatenating genome tag with seqnames tag.

Author(s)

VJ Carey <stvjc@channing.harvard.edu>

Examples

```
# do a p-value truncation if you want to reduce compression
## Not run: # ggbio July 2018
data(ebicat_2020_04_30)
library(GenomeInfoDb)
seqlevelsStyle(ebicat_2020_04_30) = "UCSC"
traitsManh(ebicat_2020_04_30)

## End(Not run)
```

[,gwaswloc,ANY,ANY,ANY-method
extractor for gwaswloc

Description

extractor for gwaswloc

Usage

```
## S4 method for signature 'gwaswloc,ANY,ANY,ANY'  
x[i, j, ..., drop = FALSE]
```

Arguments

| | |
|------|---------------|
| x | gwaswloc |
| i | index |
| j | index |
| ... | addtl indices |
| drop | logical(1) |

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