

Package ‘SAIGEgds’

March 30, 2021

Type Package

Title Scalable Implementation of Generalized mixed models using GDS files in Phenome-Wide Association Studies

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Depends R (>= 3.5.0), gdsfmt (>= 1.20.0), SeqArray (>= 1.24.1), Rcpp

LinkingTo Rcpp, RcppArmadillo, RcppParallel

Imports methods, stats, utils, RcppParallel, SPAtest (>= 3.0.0)

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Description Scalable implementation of generalized mixed models with highly optimized C++ implementation and integration with Genomic Data Structure (GDS) files. It is designed for single variant tests in large-scale phenome-wide association studies (PheWAS) with millions of variants and samples, controlling for sample structure and case-control imbalance. The implementation is based on the original SAIGE R package (v0.29.4.4). SAIGEgds also implements some of the SPAtest functions in C to speed up the calculation of Saddlepoint approximation. Benchmarks show that SAIGEgds is 5 to 6 times faster than the original SAIGE R package.

License GPL-3

SystemRequirements C++11, GNU make

VignetteBuilder knitr

ByteCompile TRUE

URL <https://github.com/AbbVie-ComputationalGenomics/SAIGEgds>

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R topics documented:

SAIGEgds-package	2
pACAT	3
seqAssocGLMM_SPA	4
seqAssocGLMM_spaACAT_O	6
seqAssocGLMM_spaACAT_V	8
seqAssocGLMM_spaBurden	10
seqFitNullGLMM_SPA	12
seqSAIGE_LoadPval	14

Index	16
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SAIGEgds-package	<i>Scalable Implementation of Generalized mixed models in Phenome-Wide Association Studies using GDS files</i>
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Description

Scalable and accurate implementation of generalized mixed mode with the support of Genomic Data Structure (GDS) files and highly optimized C++ implementation. It is designed for single variant tests in large-scale phenome-wide association studies (PheWAS) with millions of variants and hundreds of thousands of samples, e.g., UK Biobank genotype data, controlling for case-control imbalance and sample structure in single variant association studies.

The implementation of SAIGEgds is based on the original SAIGE R package (v0.29.4.4) [Zhou et al. 2018] <https://github.com/weizhouUMICH/SAIGE/releases/tag/v0.29.4.4>. All of the calculation with single-precision floating-point numbers in SAIGE are replaced by the double-precision calculation in SAIGEgds. SAIGEgds also implements some of the SPAtest functions in C to speed up the calculation of Saddlepoint Approximation.

Details

Package: SAIGEgds
 Type: Package
 License: GPL version 3

Author(s)

Xiuwen Zheng <xiuwen.zheng@abbvie.com>, Wei Zhou (the original author of the SAIGE R package, <https://github.com/weizhouUMICH/SAIGE>)

References

Zheng X, Davis J, Wade. SAIGEgds – an efficient statistical tool for large-scale PheWAS with mixed models. *Bioinformatics* (2020). DOI: 10.1093/bioinformatics/btaa731.

Zhou W, Nielsen JB, Fritsche LG, Dey R, Gabrielsen ME, Wolford BN, LeFaive J, VandeHaar P, Gagliano SA, Gifford A, Bastarache LA, Wei WQ, Denny JC, Lin M, Hveem K, Kang HM, Abeca-

sis GR, Willer CJ, Lee S. Efficiently controlling for case-control imbalance and sample relatedness in large-scale genetic association studies. *Nat Genet* (2018). Sep;50(9):1335-1341.

Zheng X, Gogarten S, Lawrence M, Stilp A, Conomos M, Weir BS, Laurie C, Levine D. SeqArray – A storage-efficient high-performance data format for WGS variant calls. *Bioinformatics* (2017). DOI: 10.1093/bioinformatics/btx145.

Examples

```
# open the GDS file
fn <- system.file("extdata", "grm1k_10k_snp.gds", package="SAIGEgds")
gdsfile <- seqOpen(fn)

# load phenotype
phenofn <- system.file("extdata", "pheno.txt.gz", package="SAIGEgds")
pheno <- read.table(phenofn, header=TRUE, as.is=TRUE)
head(pheno)

# fit the null model
glmm <- seqFitNullGLMM_SPA(y ~ x1 + x2, pheno, gdsfile, trait.type="binary")

# p-value calculation
assoc <- seqAssocGLMM_SPA(gdsfile, glmm, mac=10)

head(assoc)

# close the GDS file
seqClose(gdsfile)
```

pACAT

Cauchy Combination Test

Description

P-value calculation from Cauchy combination test.

Usage

```
pACAT(p, w=NULL)
pACAT2(p, maf, wbeta=c(1,25))
```

Arguments

p	a numeric vector for p-values
w	weight for each p-value
maf	minor allele frequency for each p-value
wbeta	weights for per-variant effect, using beta distribution <code>dbeta()</code> according to variant's MAF

Value

Return a single number for the combined p-value.

References

Liu Y., Cheng S., Li Z., Morrison A.C., Boerwinkle E., Lin X.; ACAT: A Fast and Powerful p Value Combination Method for Rare-Variant Analysis in Sequencing Studies. *Am J Hum Genetics* 104, 410-421 (2019).

See Also

[seqFitNullGLMM_SPA](#), [seqAssocGLMM_SPA](#)

Examples

```
p1 <- 10^-4
p2 <- 10^-5
p3 <- 10^-(3:20)
sapply(p3, function(p) pACAT(c(p1, p2, p)))

pACAT2(c(10^-4, 10^-6), c(0.01, 0.005))
```

seqAssocGLMM_SPA	<i>P-value calculation</i>
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Description

P-value calculations using variance approximation and an adjustment of Saddlepoint approximation.

Usage

```
seqAssocGLMM_SPA(gdsfile, modobj, maf=NaN, mac=10, missing=0.1, dsnode="",
  spa.pval=0.05, var.ratio=NaN, res.savefn="", res.compress="LZMA",
  parallel=FALSE, verbose=TRUE)
```

Arguments

gdsfile	a SeqArray GDS filename, or a GDS object
modobj	an R object for SAIGE model parameters
maf	minor allele frequency threshold (checking \geq maf), NaN for no filter
mac	minor allele count threshold (checking \geq mac), NaN for no filter
missing	missing threshold for variants (checking \leq missing), NaN for no filter
dsnode	"" for automatically searching the GDS nodes "genotype" and "annotation/format/DS", or use a user-defined GDS node in the file
spa.pval	the p-value threshold for SPA adjustment, 0.05 by default (since normal approximation performs well when the test statistic is close to the mean)
var.ratio	NaN for using the estimated variance ratio in the model fitting, or a user-defined variance ratio
res.savefn	an RData or GDS file name, "" for no saving
res.compress	the compression method for the output file, it should be one of LZMA, LZMA_RA, ZIP, ZIP_RA and none

parallel	FALSE (serial processing), TRUE (multicore processing), a numeric value for the number of cores, or other value; parallel is passed to the argument <code>cl</code> in seqParallel , see seqParallel for more details
verbose	if TRUE, show information

Details

The original SAIGE R package uses 0.05 as a threshold for unadjusted p-values (based on asymptotic normality) to further calculate adjusted p-values (Saddlepoint approximation, SPA). If `var.ratio=NaN`, the average of variance ratios (`mean(modobj$var.ratio$ratio)`) is used instead. For more details of SAIGE algorithm, please refer to the SAIGE paper [Zhou et al. 2018] (see the reference section).

Value

Return a `data.frame` with the following components if not saving to a file:

<code>id</code>	variant ID in the GDS file;
<code>chr</code>	chromosome;
<code>pos</code>	position;
<code>rs.id</code>	the RS IDs if it is available in the GDS file;
<code>ref</code>	the reference allele;
<code>alt</code>	the alternative allele;
<code>AF.alt</code>	allele frequency for the alternative allele; the minor allele frequency is $\min(\text{AF.alt}, 1-\text{AF.alt})$;
<code>mac</code>	minor allele count; the allele count for the alternative allele is $\text{ifelse}(\text{AF.alt} \leq 0.5, \text{mac}, 2 * \text{num} - \text{mac})$;
<code>num</code>	the number of samples with non-missing genotypes;
<code>beta</code>	beta coefficient, odds ratio if binary outcomes (alternative allele vs. reference allele);
<code>SE</code>	standard error for beta coefficient;
<code>pval</code>	adjusted p-value with the Saddlepoint approximation method;
<code>p.norm</code>	p-values based on asymptotic normality (could be 0 if it is too small, e.g., $\text{pnorm}(-50) = 0$ in R; used for checking only)
<code>converged</code>	whether the SPA algorithm converges or not for adjusted p-values.

Author(s)

Xiuwen Zheng

References

Zhou W, Nielsen JB, Fritsche LG, Dey R, Gabrielsen ME, Wolford BN, LeFaive J, VandeHaar P, Gagliano SA, Gifford A, Bastarache LA, Wei WQ, Denny JC, Lin M, Hveem K, Kang HM, Abecasis GR, Willer CJ, Lee S. Efficiently controlling for case-control imbalance and sample relatedness in large-scale genetic association studies. *Nat Genet* (2018). Sep;50(9):1335-1341.

See Also

[seqAssocGLMM_SPA](#), [seqSAIGE_LoadPval](#)

Examples

```
# open a GDS file
fn <- system.file("extdata", "grm1k_10k_snp.gds", package="SAIGEgds")
gdsfile <- seqOpen(fn)

# load phenotype
phenofn <- system.file("extdata", "pheno.txt.gz", package="SAIGEgds")
pheno <- read.table(phenofn, header=TRUE, as.is=TRUE)
head(pheno)

# fit the null model
glmm <- seqFitNullGLMM_SPA(y ~ x1 + x2, pheno, gdsfile, trait.type="binary")

# p-value calculation
assoc <- seqAssocGLMM_SPA(gdsfile, glmm, mac=10)
head(assoc)

# close the GDS file
seqClose(gdsfile)
```

```
seqAssocGLMM_spaACAT_O
```

ACAT-V tests

Description

ACAT-O combined p-value calculations using mixed models and the Saddlepoint approximation method for case-control imbalance.

Usage

```
seqAssocGLMM_spaACAT_O(gdsfile, modobj, units, wbeta=AggrParamBeta,
  burden.mac=10, burden.summac=3, dsnode="", spa.pval=0.05, var.ratio=NaN,
  res.savefn="", res.compress="LZMA", parallel=FALSE,
  verbose=TRUE, verbose.maf=TRUE)
```

Arguments

<code>gdsfile</code>	a SeqArray GDS filename, or a GDS object
<code>modobj</code>	an R object for SAIGE model parameters
<code>units</code>	a list of units of selected variants, with S3 class "SeqUnitListClass" defined in the SeqArray package
<code>wbeta</code>	weights for per-variant effect, using beta distribution <code>dbeta()</code> according to variant's MAF; a length-two vector, or a matrix with two rows for multiple beta parameters; by default, using <code>beta(1,1)</code> and <code>beta(1,25)</code> both
<code>burden.mac</code>	a threshold of minor allele count for using burden test instead of single variant test if <code>mac < burden.mac</code>
<code>burden.summac</code>	a threshold for the weighted sum of minor allele counts in burden test (checking <code>>= burden.summac</code>)
<code>dsnode</code>	"" for automatically searching the GDS nodes "genotype" and "annotation/format/DS", or use a user-defined GDS node in the file

spa.pval	the p-value threshold for SPA adjustment, 0.05 by default
var.ratio	NaN for using the estimated variance ratio in the model fitting, or a user-defined variance ratio
res.savefn	an RData or GDS file name, "" for no saving
res.compress	the compression method for the output file, it should be one of LZMA, LZMA_RA, ZIP, ZIP_RA and none
parallel	FALSE (serial processing), TRUE (multicore processing), a numeric value for the number of cores, or other value; parallel is passed to the argument cl in seqParallel , see seqParallel for more details
verbose	if TRUE, show information
verbose.maf	if TRUE, show summary of MAFs in units

Details

The original SAIGE R package uses 0.05 as a threshold for unadjusted p-values to further calculate SPA-adjusted p-values. If `var.ratio=NaN`, the average of variance ratios (`mean(modobj$var.ratio$ratio)`) is used instead. For more details of SAIGE algorithm, please refer to the SAIGE paper [Zhou et al. 2018] (see the reference section). No SKAT implementation.

Value

Return a data.frame with the following components if not saving to a file: chr, chromosome; start, a starting position; end, an ending position; numvar, the number of variants in a window; summac, the weighted sum of minor allele counts; beta, beta coefficient, odds ratio if binary outcomes; SE, standard error for beta coefficient; pval, adjusted p-value with Saddlepoint approximation;

p.norm p-values based on asymptotic normality (could be 0 if it is too small, e.g., `pnorm(-50) = 0` in R; used for checking only

cvg, whether the SPA algorithm converges or not for adjusted p-value.

Author(s)

Xiuwen Zheng

References

Liu Y., Chen S., Li Z., Morrison A.C., Boerwinkle E., Lin X. ACAT: A Fast and Powerful p Value Combination Method for Rare-Variant Analysis in Sequencing Studies. *Am J Hum Genetics* 104, 410-421 (2019).

See Also

[seqAssocGLMM_spaBurden](#), [seqAssocGLMM_spaACAT_V](#)

Examples

```
# open a GDS file
fn <- system.file("extdata", "grm1k_10k_snp.gds", package="SAIGEgds")
gdsfile <- seqOpen(fn)

# load phenotype
```

```

phenofn <- system.file("extdata", "pheno.txt.gz", package="SAIGEgds")
pheno <- read.table(phenofn, header=TRUE, as.is=TRUE)
head(pheno)

# fit the null model
glmm <- seqFitNullGLMM_SPA(y ~ x1 + x2, pheno, gdsfile, trait.type="binary")

# get a list of variant units for burden tests
units <- seqUnitSlidingWindows(gdsfile, win.size=500, win.shift=250)

assoc <- seqAssocGLMM_spaACAT_0(gdsfile, glmm, units)
head(assoc)

# close the GDS file
seqClose(gdsfile)

```

```
seqAssocGLMM_spaACAT_V
```

ACAT-V tests

Description

ACAT-V p-value calculations using mixed models and the Saddlepoint approximation method for case-control imbalance.

Usage

```
seqAssocGLMM_spaACAT_V(gdsfile, modobj, units, wbeta=AggrParamBeta,
  burden.mac=10, burden.summac=3, dsnode="", spa.pval=0.05, var.ratio=NaN,
  res.savefn="", res.compress="LZMA", parallel=FALSE,
  verbose=TRUE, verbose.maf=TRUE)
```

Arguments

<code>gdsfile</code>	a SeqArray GDS filename, or a GDS object
<code>modobj</code>	an R object for SAIGE model parameters
<code>units</code>	a list of units of selected variants, with S3 class "SeqUnitListClass" defined in the SeqArray package
<code>wbeta</code>	weights for per-variant effect, using beta distribution <code>dbeta()</code> according to variant's MAF; a length-two vector, or a matrix with two rows for multiple beta parameters; by default, using <code>beta(1,1)</code> and <code>beta(1,25)</code> both
<code>burden.mac</code>	a threshold of minor allele count for using burden test instead of single variant test if <code>mac < burden.mac</code>
<code>burden.summac</code>	a threshold for the weighted sum of minor allele counts in burden test (checking <code>>= burden.summac</code>)
<code>dsnode</code>	"" for automatically searching the GDS nodes "genotype" and "annotation/format/DS", or use a user-defined GDS node in the file
<code>spa.pval</code>	the p-value threshold for SPA adjustment, 0.05 by default
<code>var.ratio</code>	NaN for using the estimated variance ratio in the model fitting, or a user-defined variance ratio

res.savefn	an RData or GDS file name, "" for no saving
res.compress	the compression method for the output file, it should be one of LZMA, LZMA_RA, ZIP, ZIP_RA and none
parallel	FALSE (serial processing), TRUE (multicore processing), a numeric value for the number of cores, or other value; parallel is passed to the argument cl in seqParallel , see seqParallel for more details
verbose	if TRUE, show information
verbose.maf	if TRUE, show summary of MAFs in units

Details

Liu Y., Chen S., Li Z., Morrison A.C., Boerwinkle E., Lin X. ACAT: A Fast and Powerful p Value Combination Method for Rare-Variant Analysis in Sequencing Studies. *Am J Hum Genetics* 104, 410-421 (2019).

Value

Return a data.frame with the following components if not saving to a file: chr, chromosome; start, a starting position; end, an ending position; numvar, the number of variants in a window; summac, the weighted sum of minor allele counts; beta, beta coefficient, odds ratio if binary outcomes); SE, standard error for beta coefficient; pval, adjusted p-value with Saddlepoint approximation;

p.norm p-values based on asymptotic normality (could be 0 if it is too small, e.g., $\text{pnorm}(-50) = 0$ in R; used for checking only

cvg, whether the SPA algorithm converges or not for adjusted p-value.

Author(s)

Xiuwen Zheng

References

XX

See Also

[seqAssocGLMM_spaBurden](#), [seqAssocGLMM_spaACAT_0](#)

Examples

```
# open a GDS file
fn <- system.file("extdata", "grm1k_10k_snp.gds", package="SAIGEgds")
gdsfile <- seqOpen(fn)

# load phenotype
phenofn <- system.file("extdata", "pheno.txt.gz", package="SAIGEgds")
pheno <- read.table(phenofn, header=TRUE, as.is=TRUE)
head(pheno)

# fit the null model
glimm <- seqFitNullGLMM_SPA(y ~ x1 + x2, pheno, gdsfile, trait.type="binary")
```

```
# get a list of variant units for burden tests
units <- seqUnitSlidingWindows(gdsfile, win.size=500, win.shift=250)

assoc <- seqAssocGLMM_spaACAT_V(gdsfile, glmm, units)
head(assoc)

# close the GDS file
seqClose(gdsfile)
```

```
seqAssocGLMM_spaBurden
```

Burden tests

Description

Burden p-value calculations using mixed models and the Saddlepoint approximation method for case-control imbalance.

Usage

```
seqAssocGLMM_spaBurden(gdsfile, modobj, units, wbeta=AggrParamBeta,
  summac=3, dsnode="", spa.pval=0.05, var.ratio=NaN, res.savefn="",
  res.compress="LZMA", parallel=FALSE, verbose=TRUE, verbose.maf=TRUE)
```

Arguments

<code>gdsfile</code>	a SeqArray GDS filename, or a GDS object
<code>modobj</code>	an R object for SAIGE model parameters
<code>units</code>	a list of units of selected variants, with S3 class "SeqUnitListClass" defined in the SeqArray package
<code>wbeta</code>	weights for per-variant effect, using beta distribution <code>dbeta()</code> according to variant's MAF; a length-two vector, or a matrix with two rows for multiple beta parameters; by default, using <code>beta(1,1)</code> and <code>beta(1,25)</code> both
<code>summac</code>	a threshold for the weighted sum of minor allele counts (checking \geq <code>summac</code>)
<code>dsnode</code>	"" for automatically searching the GDS nodes "genotype" and "annotation/format/DS", or use a user-defined GDS node in the file
<code>spa.pval</code>	the p-value threshold for SPA adjustment, 0.05 by default
<code>var.ratio</code>	NaN for using the estimated variance ratio in the model fitting, or a user-defined variance ratio
<code>res.savefn</code>	an RData or GDS file name, "" for no saving
<code>res.compress</code>	the compression method for the output file, it should be one of LZMA, LZMA_RA, ZIP, ZIP_RA and none
<code>parallel</code>	FALSE (serial processing), TRUE (multicore processing), a numeric value for the number of cores, or other value; <code>parallel</code> is passed to the argument <code>c1</code> in seqParallel , see seqParallel for more details
<code>verbose</code>	if TRUE, show information
<code>verbose.maf</code>	if TRUE, show summary of MAFs in units

Details

The original SAIGE R package uses 0.05 as a threshold for unadjusted p-values to further calculate SPA-adjusted p-values. If `var.ratio=NaN`, the average of variance ratios (`mean(modobj$var.ratio$ratio)`) is used instead. For more details of SAIGE algorithm, please refer to the SAIGE paper [Zhou et al. 2018] (see the reference section).

Value

Return a `data.frame` with the following components if not saving to a file: `chr`, chromosome; `start`, a starting position; `end`, an ending position; `numvar`, the number of variants in a window; `summac`, the weighted sum of minor allele counts; `beta`, beta coefficient, odds ratio if binary outcomes; `SE`, standard error for beta coefficient; `pval`, adjusted p-value with Saddlepoint approximation;

`p.norm` p-values based on asymptotic normality (could be 0 if it is too small, e.g., `pnorm(-50) = 0` in R; used for checking only)

`cvg`, whether the SPA algorithm converges or not for adjusted p-value.

Author(s)

Xiuwen Zheng

References

Zhou W, Nielsen JB, Fritsche LG, Dey R, Gabrielsen ME, Wolford BN, LeFaive J, VandeHaar P, Gagliano SA, Gifford A, Bastarache LA, Wei WQ, Denny JC, Lin M, Hveem K, Kang HM, Abecasis GR, Willer CJ, Lee S. Efficiently controlling for case-control imbalance and sample relatedness in large-scale genetic association studies. *Nat Genet* (2018). Sep;50(9):1335-1341.

See Also

[seqAssocGLMM_spaACAT_V](#), [seqAssocGLMM_spaACAT_0](#)

Examples

```
# open a GDS file
fn <- system.file("extdata", "grm1k_10k_snp.gds", package="SAIGEgds")
gdsfile <- seqOpen(fn)

# load phenotype
phenofn <- system.file("extdata", "pheno.txt.gz", package="SAIGEgds")
pheno <- read.table(phenofn, header=TRUE, as.is=TRUE)
head(pheno)

# fit the null model
glmm <- seqFitNullGLMM_SPA(y ~ x1 + x2, pheno, gdsfile, trait.type="binary")

# get a list of variant units for burden tests
units <- seqUnitSlidingWindows(gdsfile, win.size=500, win.shift=250)

assoc <- seqAssocGLMM_spaBurden(gdsfile, glmm, units)
head(assoc)

# close the GDS file
seqClose(gdsfile)
```

seqFitNullGLMM_SPA *Fit the null model with GRM*

Description

Fit the null model in the mixed model framework with genetic relationship matrix (GRM).

Usage

```
seqFitNullGLMM_SPA(formula, data, gdsfile, trait.type=c("binary", "quantitative"),
  sample.col="sample.id", maf=0.005, missing.rate=0.01, max.num.snp=1000000L,
  variant.id=NULL, inv.norm=TRUE, X.transform=TRUE, tol=0.02, maxiter=20L,
  nrun=30L, tolPCG=1e-5, maxiterPCG=500L, num.marker=30L, tau.init=c(0,0),
  traceCVcutoff=0.0025, ratioCVcutoff=0.001, geno.sparse=TRUE, num.thread=1L,
  model.savefn="", seed=200L, fork.loading=FALSE, verbose=TRUE)
```

Arguments

formula	an object of class formula (or one that can be coerced to that class), e.g., $y \sim x_1 + x_2$, see lm
data	a data frame for the formulas
gdsfile	a SeqArray GDS filename, or a GDS object
trait.type	"binary" for binary outcomes, "quantitative" for continuous outcomes
sample.col	the column name of sample IDs corresponding to the GDS file
maf	minor allele frequency for imported genotypes (checking \geq maf), if variant.id=NULL; NaN for no filter
missing.rate	threshold of missing rate (checking \leq missing.rate), if variant.id=NULL; NaN for no filter
max.num.snp	the maximum number of SNPs used, or -1 for no limit
variant.id	a list of variant IDs, used to construct GRM
inv.norm	if TRUE, perform inverse normal transformation on residuals for quantitative outcomes, see the reference [Sofer, 2019]
X.transform	if TRUE, perform QR decomposition on the design matrix
tol	overall tolerance for model fitting
maxiter	the maximum number of iterations for model fitting
nrun	the number of random vectors in the trace estimation
tolPCG	tolerance of PCG iterations
maxiterPCG	the maximum number of PCG iterations
num.marker	the number of SNPs used to calculate the variance ratio
tau.init	a 2-length numeric vector, the initial values for variance components, tau; for binary traits, the first element is always be set to 1. if tau.init is not specified, the second element will be 0.5 for binary traits
traceCVcutoff	the threshold for coefficient of variation (CV) for the trace estimator, and the number of runs for trace estimation will be increased until the CV is below the threshold

ratioCVcutoff	the threshold for coefficient of variation (CV) for estimating the variance ratio, and the number of randomly selected markers will be increased until the CV is below the threshold
geno.sparse	if TRUE, store the sparse structure for genotypes; otherwise, save genotypes in a 2-bit dense matrix; see details
num.thread	the number of threads
model.savefn	the filename of model output, R data file '.rda', '.RData', or '.rds'
seed	an integer as a seed for random numbers
fork.loading	load genotypes via forking or not; forking processes in Unix can reduce loading time of genotypes, but may double the memory usage; not applicable on Windows
verbose	if TRUE, show information

Details

Utilizing the sparse structure of genotypes could significantly improve the computational efficiency of model fitting, but it also increases the memory usage. For more details of SAIGE algorithm, please refer to the SAIGE paper [Zhou et al. 2018] (see the reference section).

Value

Returns a list with the following components:

coefficients	the beta coefficients for fixed effects;
tau	a numeric vector of variance components 'Sigma_E' and 'Sigma_G';
linear.predictors	the linear fit on link scale;
fitted.values	fitted values from objects returned by modeling functions using <code>glm.fit</code> ;
residuals	residuals;
cov	covariance matrix of beta coefficients;
converged	whether the model is fitted or not;
obj.noK	internal use, returned object from the SPAtest package;
var.ratio	a data.frame with columns 'id' (variant.id), 'maf' (minor allele frequency), 'mac' (minor allele count), 'var1' (the variance of score statistic), 'var2' (a variance estimate without accounting for estimated random effects) and 'ratio' (var1/var2, estimated variance ratio for variance approximation);
trait.type	either "binary" or "quantitative";
sample.id	the sample IDs used in the model fitting;
variant.id	the variant IDs used in the model fitting.

Author(s)

Xiuwen Zheng

References

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See Also

[seqAssocGLMM_SPA](#)

Examples

```
# open a GDS file
fn <- system.file("extdata", "grm1k_10k_snp.gds", package="SAIGEgds")
gdsfile <- seqOpen(fn)

# load phenotype
phenofn <- system.file("extdata", "pheno.txt.gz", package="SAIGEgds")
pheno <- read.table(phenofn, header=TRUE, as.is=TRUE)
head(pheno)

# fit the null model
glm <- seqFitNullGLMM_SPA(y ~ x1 + x2, pheno, gdsfile, trait.type="binary")
glm

# close the GDS file
seqClose(gdsfile)
```

seqSAIGE_LoadPval *Load the association results*

Description

Load the association results from an RData, RDS or GDS file.

Usage

```
seqSAIGE_LoadPval(fn, varnm=NULL, index=NULL, verbose=TRUE)
```

Arguments

fn	RData, RDS or GDS file names, merging datasets if multiple files
varnm	NULL, or a character vector to include the column names; e.g., c("chr", "position", "rs.id", "ref")
index	NULL, or a logical/numeric vector for a set of rows
verbose	if TRUE, show information

Value

Return a data.frame including p-values.

Author(s)

Xiuwen Zheng

See Also

[seqFitNullGLMM_SPA](#), [seqAssocGLMM_SPA](#)

Examples

```
(fn <- system.file("unitTests", "saige_pval.rds", package="SAIGEgds"))  
pval <- seqSAIGE_LoadPval(fn)
```

```
names(pval)  
# [1] "id"      "chr"      "pos"      "rs.id"    "ref"  
# [6] "alt"     "AF.alt"   "AC.alt"   "num"      "beta"  
# [11] "SE"      "pval"     "pval.noadj" "converged"
```

```
head(pval)
```

Index

* Cauchy

pACAT, 3

* GDS

SAIGEgds-package, 2

seqAssocGLMM_SPA, 4

seqAssocGLMM_spaACAT_0, 6

seqAssocGLMM_spaACAT_V, 8

seqAssocGLMM_spaBurden, 10

seqFitNullGLMM_SPA, 12

seqSAIGE_LoadPval, 14

* association

pACAT, 3

SAIGEgds-package, 2

seqAssocGLMM_SPA, 4

seqAssocGLMM_spaACAT_0, 6

seqAssocGLMM_spaACAT_V, 8

seqAssocGLMM_spaBurden, 10

seqFitNullGLMM_SPA, 12

seqSAIGE_LoadPval, 14

* genetics

SAIGEgds-package, 2

seqAssocGLMM_SPA, 4

seqAssocGLMM_spaACAT_0, 6

seqAssocGLMM_spaACAT_V, 8

seqAssocGLMM_spaBurden, 10

seqFitNullGLMM_SPA, 12

seqSAIGE_LoadPval, 14

lm, 12

pACAT, 3

pACAT2 (pACAT), 3

SAIGEgds (SAIGEgds-package), 2

SAIGEgds-package, 2

seqAssocGLMM_SPA, 4, 4, 5, 14, 15

seqAssocGLMM_spaACAT_0, 6, 9, 11

seqAssocGLMM_spaACAT_V, 7, 8, 11

seqAssocGLMM_spaBurden, 7, 9, 10

seqFitNullGLMM_SPA, 4, 12, 15

seqParallel, 5, 7, 9, 10

seqSAIGE_LoadPval, 5, 14