# Package 'Gimpute'

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<b>Title</b> Gimpute: An efficient genetic data processing and imputation pipeline	
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<b>Description</b> Genotype imputation is essential for genome-wide association studies (GWAS) to retrieve information of untyped variants and facilitate comparability across studies. Based on widely used and freely available tools, we have developed Gimpute, an automated processing and imputation pipeline for genome-wide association data. Gimpute includes processing steps for genotype liftOver, quality control, population outlier detection, haplotype pre-phasing, imputation, post imputation, data management and the extension to other existing pipeline.	
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checkAlign2ref

Check the alignment with the imputation reference panel

# Description

Perform the alignment against a reference panel by considering the following parameters: variant name, genomic position and the allele profile. Output files are generated sequentially depending on their previous PLINK data.

# Usage

checkAlign2ref(plink, inputPrefix, referencePanel, bimReferenceFile, out2,
 out2.snp, out3, out3.snp, out4, out4.snp, out4.snpRetained, nCore = 25)

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### **Arguments**

plink	an executable program in either the current working directory or somewhere in the command path.	
inputPrefix	the prefix of the input PLINK files.	
referencePan	el	
	a string indicating the type of imputation reference panels is used: $c("1000Gphase1v3\_macGT1", "1000Gphase3")$ .	
bimReference	File	
	the reference file used for the alignment, which is a PLINK BIM alike format file.	
out2	the prefix of the output PLINK binary files after removing SNPs whose genomic positions are not in the imputation reference, taking SNP names into account.	
out2.snp	the output plain text file that stores the removed SNPs whose genomic positions are not in the imputation reference, taking SNP names into account.	
out3	the prefix of the output PLINK binary files after removing SNPs whose genomic positions are not in the imputation reference, ingoring SNP names.	
out3.snp	the output plain text file that stores the removed SNPs whose genomic positions are not in the imputation reference, ingoring SNP names.	
out4	the prefix of the output PLINK binary files after removing SNPs whose alleles are not in the imputation reference, taking their genomic positions into account.	
out4.snp	the output plain text file that stores the removed SNPs whose alleles are not in the imputation reference, taking their genomic positions into account.	
out4.snpRetained		
	the output plain text file that stores the removed SNPs whose alleles are in the imputation reference, taking their genomic positions into account.	

# **Details**

nCore

The output files are genrated in order. Genomic position includes chromosomal location and base-pair position of the individual variant. All monomorphic SNPs are retained for further processing.

the number of cores used for computation. This can be tuned along with nThread.

### Value

The set of aligned PLINK files from your own study compared with the imputation reference.

# Author(s)

Junfang Chen

4 chrWiseSplit

chrWiseSplit Split genome-wide genotyping data into chromosome-wide PLINK binary files.	-
---	---

### **Description**

Split the whole genome genotyping data chromosome-wise; allow parallel computating for all chromosomes

#### Usage

```
chrWiseSplit(plink, inputPrefix, chrXPAR1suffix, chrXPAR2suffix, nCore = 25)
```

### **Arguments**

plink an executable program in either the current working directory or somewhere in the command path.

inputPrefix the prefix of the input PLINK binary files before splitting.

chrXPAR1suffix

if chromosome 25 is available and with PAR1, then generate the suffix with X\_PAR1 for chrX\_PAR1.

chrXPAR2suffix

if chromosome 25 is available and with PAR2, then generate the suffix with X\_PAR2 for chrX\_PAR2.

ncore the number of cores used for parallel computation. The default value is 25.

#### Details

If chromosome 25 is also available, namely the pseudo-autosomal region of chromosome X, then further split chr25 (PAR or Chr\_XY) into PAR1 and PAR2 according to the genomic coordination GRCh37 from https://en.wikipedia.org/wiki/Pseudoautosomal\_region. The locations of the PARs within GRCh37 are: PAR1 X 60001 2699520; PAR2 X 154931044 155260560.

#### Value

The output PLINK binary files for each chromosome with the same prefix as the inputPrefix but appended with the chromosome codes, and possibly also the logical value for the pseudo-autosomal region (PAR) indicating if PAR exists in the input genotyping data or not.

# Author(s)

Junfang Chen

```
## In the current working directory
bedFile <- system.file("extdata", "alignedData.bed", package="Gimpute")
bimFile <- system.file("extdata", "alignedData.bim", package="Gimpute")
famFile <- system.file("extdata", "alignedData.fam", package="Gimpute")
system(paste0("scp ", bedFile, " ."))</pre>
```

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```
system(paste0("scp ", bimFile, " ."))
system(paste0("scp ", famFile, " ."))
inputPrefix <- "alignedData"
chrXPAR1suffix <- "X_PAR1"
chrXPAR2suffix <- "X_PAR2"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## chrWiseSplit(plink, inputPrefix, chrXPAR1suffix, chrXPAR2suffix, nCore)</pre>
```

chunk4eachChr

Chunk each chromosome into multiple segments

## **Description**

Chunk each chromosome genotyping data into multiple segments by a predefined window size.

#### Usage

```
chunk4eachChr(inputPrefix, outputPrefix, chrs, windowSize = 3e+06)
```

# **Arguments**

```
inputPrefix the prefix of the input PLINK .bim file for each chromosome, without the chromosome codes.
outputPrefix the prefix of the output pure text files that keep all the chunks for each chromosome separately.
chrs specify the chromosome codes for chunking.
windowSize the window size of each chunk. The default value is 3000000.
```

### Value

The output pure text files include all the chunks for each chromosome separately.

#### Author(s)

Junfang Chen

```
## In the current working directory
bimFile <- system.file("extdata", "gwas_data_chr23.bim", package="Gimpute")
system(paste0("scp ", bimFile, " ."))
inputPrefix <- "gwas_data_chr"
outputPrefix <- "chunks_chr"
chrs <- 23
print(chrs)
chunk4eachChr(inputPrefix, outputPrefix, chrs, windowSize=3000000)</pre>
```

```
computeInfoByQctool
```

Calculate the info score by QCTOOL

# Description

Calculate the info score by QCTOOL for a set of SNPs from .GEN files.

### Usage

```
computeInfoByQctool(qctool, inputSuffix, outputInfoFile)
```

### **Arguments**

an executable program in either the current working directory or somewhere in

the command path.

inputSuffix the suffix of input .GEN files within current directory.

outputInfoFile

the output info scores file consisting of two columns: all SNPs from .GEN files

and their info scores.

### **Details**

These .GEN files may come from the output of impute4 results. The intermediate generated files are retained in the directory.

# Value

A pure text file contains the info scores of all SNPs from .GEN files with two columns: SNP names and the corresponding info scores.

### Author(s)

Junfang Chen

```
## In the current working directory
inputSuffix <- "noINDEL.gen"
outputInfoFile <- "infoScore.txt"
## computeInfoByQctool(qctool, inputSuffix, outputInfoFile)</pre>
```

extractByGenipe 7

# Description

Extract imputed markers located in a specific genomic region using Genipe. Note that, 1.) 'bed' PLINK binary format is specifically used for the output format. 2.) The markers of the whole chromosome are extracted together. For the filtering of maf and info will be done during post imputation.

### Usage

```
extractByGenipe(inputImpute2, inputMAP, outputPrefix, format, prob)
```

### **Arguments**

inputImpute2 the output from IMPUTE2.

inputMAP the output PLINK MAP file from Genipe, which will be used for generating

markers in a text file (only one column without column name).

outputPrefix the prefix of the output files. [impute2\_extractor]

format the output format. Can specify either "impute2" for probabilities (same as im-

pute2 format, i.e. 3 values per sample), "dosage" for dosage values (one value between 0 and 2 by sample), "calls" for hard calls, or "bed" for Plink binary

format (with hard calls). [impute2]

prob the probability threshold used when creating a file in the dosage or call format.

[0.9]

### Value

The extracted imputed files using genipe.

### Author(s)

Junfang Chen

### References

Lemieux Perreault, L. P., et al. (2016). genipe: an automated genome-wide imputation pipeline with automatic reporting and statistical tools. Bioinformatics, 32(23), 3661-3663.

genoQC

genoQC

Quality control for genotype data

#### **Description**

Perform quality control on the genotype data.

### Usage

```
genoQC(plink, inputPrefix, snpMissCutOffpre = 0.05, sampleMissCutOff = 0.02,
   Fhet = 0.2, cutoffSubject, cutoffSNP, snpMissCutOffpost = 0.02,
   snpMissDifCutOff = 0.02, femaleChrXmissCutoff = 0.05,
   pval4autoCtl = 1e-06, pval4femaleXctl = 1e-06, outputPrefix,
   keepInterFile = TRUE)
```

### Arguments

plink

an executable program in either the current working directory or somewhere in the command path.

inputPrefix the prefix of the input PLINK binary files.

snpMissCutOffpre

the cutoff of the missingness for removing SNPs before subject removal. The default is 0.05.

sampleMissCutOff

the cutoff of the missingness for removing subjects/instances. The default is 0.02.

Fhet

the cutoff of the autosomal heterozygosity deviation. The default is 0.2.

cutoffSubject

the cutoff determines that families (subjects) with more than the predefined cutoff of Mendel errors by considering all SNPs will be removed. The default is 0.05.

cutoffSNP

the cutoff indicates that SNPs with more than the predefined cutoff of Mendel error rate will be excluded (i.e. based on the number of trios/duos). The default is 0.1.

snpMissCutOffpost

the cutoff of the missingness for removing SNPs after subject removal. The default is 0.02.

snpMissDifCutOff

the cutoff of the difference in missingness between cases and controls. The default is 0.02.

femaleChrXmissCutoff

the cutoff of the missingness in female chromosome X SNPs. The default is 0.05.

pval4autoCtl the p-value cutoff for controlling HWE test in either control or case subjects. Only autosomal SNPs are considered. The default is 0.000001

pval4femaleXctl

the p-value cutoff for controlling HWE test in female control subjects. Only chromosome X SNPs are considered. The default is 0.000001

outputPrefix the prefix of the output PLINK binary files after QC.

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keepInterFile

a logical value indicating if the intermediate processed files should be kept or not. The default is TRUE.

#### **Details**

The original PLINK files are implicitly processed by the following default steps: 1.) Set all heterozygous alleles of SNPs on male chrX as missing; 2.) SNP missingness < 0.05 (before sample removal); 3.) Subject missingness < 0.02; 4.) Remove subjects with |Fhet| >= 0.2; 5.) Reset paternal and maternal codes; 6.) SNP missingness < 0.02 (after sample removal); 7.) Remove SNPs with difference >= 0.02 of SNP missingness between cases and controls; 8.) Remove subjects or SNPs with Mendel errors for family based data. 9.) Remove chrX SNPs with missingness >= 0.05 in females. (Optional, if no chrX data); 10.) Remove autosomal SNPs with HWE p < 10-6 in controls; 11.) Remove chrX SNPs with HWE p < 10-6 in female controls. (Optional, if no chrX data).

### Value

The output PLINK binary files after QC.

# Author(s)

Junfang Chen

### References

Schizophrenia Working Group of the Psychiatric Genomics, C. (2014). Biological insights from 108 schizophrenia-associated genetic loci. Nature 511(7510): 421-427.

```
## In the current working directory
bedFile <- system.file("extdata", "genoUpdatedData.bed", package="Gimpute")</pre>
bimFile <- system.file("extdata", "genoUpdatedData.bim", package="Gimpute")</pre>
famFile <- system.file("extdata", "genoUpdatedData.fam", package="Gimpute")</pre>
system(paste0("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "genoUpdatedData"</pre>
outputPrefix <- "2_13_removedSnpHweFemaleX"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## genoQC(plink, inputPrefix,
          snpMissCutOffpre=0.05,
##
##
          sampleMissCutOff=0.02,
##
          Fhet=0.2, cutoffSubject, cutoffSNP,
##
          snpMissCutOffpost=0.02,
          snpMissDifCutOff=0.02,
##
##
          femaleChrXmissCutoff=0.05,
##
          pval4autoCtl=0.000001,
##
          pval4femaleXctl=0.000001,
##
          outputPrefix, keepInterFile=TRUE)
```

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getGroupLabel

Get the outcome label of the genotype data

### **Description**

Get the group label from the PLINK FAM file.

# Usage

```
getGroupLabel(inputFAMfile)
```

### **Arguments**

inputFAMfile the PLINK FAM file.

#### **Details**

If the input FAM file also has missing outcomes, which are shown in the sixth column of FAM file as "0", then an error is given.

### Value

The group label of the genotype data: "control" or "case" or "caseControl" indicating both groups exist.

### Author(s)

Junfang Chen

# Examples

```
famFile <- system.file("extdata", "genoUpdatedData.fam", package="Gimpute")
getGroupLabel(inputFAMfile=famFile)</pre>
```

imputedByGenipe

Impute genotypes using Genipe

# Description

Perform imputation by Genipe for the autosomal and sex chromosome prephased known haplotypes with a reference panel. Note that pre-phasing using SHAPEIT is done without the refernce haplotypes.

# Usage

```
imputedByGenipe(chrs, impRefDir, inputPrefix, shapeit, impute2, plink,
  fastaFile, segmentSize, thread4impute2, thread4shapeit)
```

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#### **Arguments**

specify the chromosomes for imputation. There are four different options ("autosomes", "1, or 2, or 3...or 22", "23", "25"). 1.) 'autosomes': will impute chromosome 1 to 22 together; 2.) 'chrs' belongs to one of (1, 2,...22) then the imputation is done just for one autosomal chromosome. 3.) '23' will do the imputation for the non-pseudoautosomal region of chromosome 23. 4.) '25'

imputes for the pseudoautosomal regions of chromosome 23.

impRefDir the directory where the imputation reference files are located.

inputPrefix the prefix of the input PLINK binary files.

shapeit an executable SHAPEIT binary program in either the current working directory

or somewhere in the command path.

impute2 an executable IMPUTE2 binary program in either the current working directory

or somewhere in the command path.

plink an executable program in either the current working directory or somewhere in

the command path.

fastaFile the human reference files for the initial strand check.

segmentSize the length of a single segment for imputation.

thread4impute2

the number of threads for the imputation.

thread4shapeit

the number of threads for phasing.

#### Value

The imputed files using genipe.

#### Author(s)

Junfang Chen

### References

Lemieux Perreault, L. P., et al. (2016). genipe: an automated genome-wide imputation pipeline with automatic reporting and statistical tools. Bioinformatics, 32(23), 3661-3663.

mergeByGenipe

Merge imputed files using Genipe

# Description

Concatenate IMPUTE2 output files and retrieve some statistics. This is automatically called by the main genipe pipeline to merge IMPUTE2 files generated for all the genomic segments. For details, see Genipe IMPUTE2 merger options.

### Usage

mergeByGenipe(inputImpute2, chr, probability, completionRate, info, outputPrefix) 12 phaseImpute

### **Arguments**

inputImpute2 the output from IMPUTE2.

chr specify the chromosome segment to be merged, on which the imputation was

made.

probability the probability threshold for no calls. [<0.9]

completionRate

the completion rate threshold for site exclusion. [<0.98]

info the measure of the observed statistical information associated with the allele

frequency estimate threshold for site exclusion. (<0.00)

outputPrefix the prefix for the imputed output files.

#### Value

The merged imputed files using genipe.

### Author(s)

Junfang Chen

#### References

Lemieux Perreault, L. P., et al. (2016). genipe: an automated genome-wide imputation pipeline with automatic reporting and statistical tools. Bioinformatics, 32(23), 3661-3663.

phaseImpute Phasing and imputation

### Description

Perform phasing, imputation and conversion from IMPUTE2 or GEN format into PLINK binary files.

#### Usage

```
phaseImpute(inputPrefix, outputPrefix, plink, shapeit, imputeTool, impute,
  qctool, gtool, windowSize = 3e+06, effectiveSize = 20000, nCore = 40,
  threshold = 0.9, outputInfoFile, referencePanel, impRefDIR, tmpImputeDir,
  keepTmpDir = TRUE)
```

# Arguments

inputPrefix the prefix of the input PLINK binary files for the imputation. outputPrefix the prefix of the output PLINK binary files after imputation.

plink an executable program in either the current working directory or somewhere in

the command path.

shapeit an executable program in either the current working directory or somewhere in

the command path.

imputeTool a string indicating the type of imputation tool is used: "impute2" or "impute4".

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impute	an executable program in either the current working directory or somewhere in the command path. It can be either "impute2" or "impute4".	
qctool	an executable program in either the current working directory or somewhere in the command path. This is only used if imputeTool is "impute4".	
gtool	an executable program in either the current working directory or somewhere in the command path.	
windowSize effectiveSiz	the window size of each chunk. The default value is 3000000.	
	this parameter controls the effective population size. Commonly denoted as Ne. A universal -Ne value of 20000 is suggested.	
nCore	the number of cores used for splitting chromosome by PLINK, phasing, imputation, genotype format modification, genotype conversion, and merging genotypes. The default value is 40.	
threshold	threshold for merging genotypes from GEN probability. Default 0.9.	
outputInfoFile		
	the output file of impute2 info scores consisting of two columns: all imputed SNPs and their info scores.	
referencePanel		
	a string indicating the type of imputation reference panels is used: " $1000Gphase1v3\_macGT1$ " or " $1000Gphase3$ ".	
impRefDIR	the directory where the imputation reference files are located.	
tmpImputeDir	the name of the temporary directory used for storing phasing and imputation results.	
keepTmpDir	a logical value indicating if the directory 'tmpImputeDir' should be kept or not. The default is TRUE.	

### **Details**

The whole imputation process mainly consists of the following steps: 1.) Phasing the input PLINK data using an existing imputation reference; 2.) Imputing the input PLINK data using phased results and an existing reference data; 3.) Converting IMPUTE2 or GEN format data into PLINK format. 4.) Combining all imputed data into whole-genome PLINK binary files. 5.) Filtering out imputed variants with bad imputation quality. Parallel computing in R is supported.

# Value

Note that chromosome X is not supported for the impute4. 1.) The filtered imputed PLINK binary files; 2.) The final PLINK binary files including bad imputed variants; 3.) A pure text file contains the info scores of all imputed SNPs with two columns: SNP names and the corresponding info scores.

#### Author(s)

Junfang Chen

#### References

- 1. Howie, B., et al. (2012). Fast and accurate genotype imputation in genome-wide association studies through pre-phasing. Nat Genet 44(8): 955-959.
- 2. Howie, B. N., et al. (2009). A flexible and accurate genotype imputation method for the next generation of genome-wide association studies. PLoS Genet 5(6): e1000529.

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3. Bycroft, C., et al. Genome-wide genetic data on~ 500,000 UK Biobank participants. BioRxiv (2017): 166298.

# **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "alignedData.bed", package="Gimpute")</pre>
bimFile <- system.file("extdata", "alignedData.bim", package="Gimpute")</pre>
famFile <- system.file("extdata", "alignedData.fam", package="Gimpute")</pre>
system(paste0("scp ", bedFile, " ."))
system(paste0("scp ", bimFile, " ."))
system(paste0("scp ", famFile, " ."))
inputPrefix <- "alignedData"</pre>
outputPrefix <- "gwasImputed"</pre>
outputInfoFile <- "infoScore.txt"</pre>
tmpImputeDir <- "tmpImpute"</pre>
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"</pre>
## phaseImpute(inputPrefix, outputPrefix,
               plink, shapeit, imputeTool, impute, qctool, gtool,
##
                windowSize=3000000, effectiveSize=20000,
##
##
               nCore=40, threshold=0.9, outputInfoFile,
##
                referencePanel, impRefDIR, tmpImputeDir, keepTmpDir=TRUE)
```

plotPCA4plink

Population outlier detection

# Description

Principle component analysis (PCA) on the genotype data is performed to detect population outliers, and the first two PCs are plotted for the visualization.

### Usage

```
plotPCA4plink(gcta, inputPrefix, nThread = 20, outputPC4subjFile,
  outputPCplotFile)
```

#### **Arguments**

gcta an executable program in either the current working directory or somewhere in the command path.

inputPrefix the prefix of the input PLINK binary files.

nThread the number of threads used for computation. The default is 20.

outputPC4subjFile

the pure text file that stores all the subject IDs and their corresponding eigenvalues of the first two principle components.

outputPCplotFile

the plot file for visualizing the first two principle components of all investigated subjects.

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### **Details**

Before population outlier detection, it's better to perform QC on the genotype data. Only autosomal genotypes are used for principle component analysis.

#### Value

The output pure text file and plot file for storing first two principle components of study subjects.

# Author(s)

Junfang Chen

### **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "QCdata.bed", package="Gimpute")
bimFile <- system.file("extdata", "QCdata.bim", package="Gimpute")
famFile <- system.file("extdata", "QCdata.fam", package="Gimpute")
system(paste0("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "QCdata"
outputPC4subjFile <- "2_13_eigenvalAfterQC.txt"
outputPCplotFile <- "2_13_eigenvalAfterQC.png" ## png format
## Not run: Requires an executable program GCTA, e.g.
## gcta <- "/home/tools/gcta64"
## plotPCA4plink(gcta, inputPrefix, nThread=20,
## outputPC4subjFile, outputPrefix)</pre>
```

postImpQC

Post imputation quality control

### **Description**

Perform quality control and data management after imputation.

# Usage

```
postImpQC(plink, inputPrefix, out1, out2, out3, out4, outputInfoFile,
  infoScore = 0.6, outputMonoSNPfile, prefixAlign2ref, missCutoff = 20,
  outRemovedSNPfile, outRetainSNPfile, referencePanel)
```

# **Arguments**

plink	an executable program in either the current working directory or somewhere in the command path.
inputPrefix	the prefix of the final imputed PLINK files.
out1	the prefix of well imputed PLINK files with the index.
out2	the prefix of well imputed PLINK files after removing any SNPs with the same positions (if any), the index is also appended.
out3	the prefix of well imputed PLINK files with the index after adding previously identified monomorphic SNPs if any.

out 4 the prefix of final well imputed PLINK files with the index.

outputInfoFile

the output file of impute2 info scores consisting of two columns: all imputed

SNPs and their info scores.

infoScore the cutoff of filtering imputation quality score for each variant. The default value

is 0.6.

outputMonoSNPfile

the output pure text file that stores the removed monomorphic SNPs, one per

line, if any.

prefixAlign2ref

the prefix of the output PLINK binary files after removing SNPs whose alleles are not in the imputation reference, taking their genomic positions into account.

missCutoff the cutoff of the least number of instances for a SNP that is not missing. The default is 20.

outRemovedSNPfile

the output file of SNPs with pre-defined missing values that are removed.

outRetainSNPfile

the output file of SNPs that are retained.

referencePanel

a string indicating the type of imputation reference panels is used: c("1000Gphase1v3\_macGT1", "1000Gphase3").

#### Value

All imputed genotype data in PLINK format, well imputed data and its variants, as well as a set of pure text files containing removed or retained SNPs.

### Author(s)

Junfang Chen

prepareAnnoFile4affy

prepare Affymetrix chip annotation file

### **Description**

Prepare Affymetrix chip annotation file into the format of interest.

#### Usage

```
prepareAnnoFile4affy(inputFile, outputFile, chipType)
```

# **Arguments**

inputFile an input pure text file that contains the chip annotation information.

outputFile an output pure text file that stores the chip annotation information in a user-

defined format.

chipType a string name defines the type of the chip annotation file: 'SNPIDstudy', and

'rsIDstudy'.

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#### **Details**

If the chip annotation file is not available for your study, it can be downloaded from http://www.well.ox.ac.uk/~wrayner/st The chip annotation file is organized into two different types:

- 1. If the snp name of your study genotype data starts with "SNP\_", then the chip type "SNPID-study" is used; Usually, Affymetrix chip data belongs to this category. The prepared output annotation file must at least consist of the following column names: SNPIDstudy, rs, chr, pos, strand.
- 2. If the snp name of your study genotype data starts with "rs", then the chip type "rsIDstudy" is used; The prepared output annotation file must at least consist of the following column names: SNPIDstudy, rs, chr, pos, strand. Illumina chip is often specified in this format.

The column "strand" must only have two kinds of values "-" and "+". Variants with all other values should be excluded.

#### Value

a pure text file that stores the prepared chip annotation information in a user-defined format.

#### Author(s)

Junfang Chen

reductExpand

Post imputation data extraction and expansion

# **Description**

Reduce well imputed dataset to have SNPs before imputation and then added genotype data that are different from the imputation reference panel.

### Usage

```
reductExpand(plink, referencePanel, inputPrefix, inputQCprefix,
    snpRefAlleleFile, snpDiffAlleleFile, snpMissPosFile, snpSameNameDifPosFile,
    reducedToSpecificfn, specificDiffAllelefn, specificMissPosfn,
    specificDiffPosfn)
```

### Arguments

plink

an executable program in either the current working directory or somewhere in the command path.

referencePanel

a string indicating the type of imputation reference panels is used: c("1000Gphase1v3\_macGT1", "1000Gphase3").

inputPrefix the prefix of final well imputed PLINK files.

inputQCprefix

the prefix of QCed PLINK files.

snpRefAlleleFile

the output plain text file that stores the removed SNPs whose alleles are in the imputation reference, taking their genomic positions into account.

18 removedDoubleProbes

snpDiffAlleleFile

the output plain text file that stores the removed SNPs whose alleles are not in the imputation reference, taking their genomic positions into account.

snpMissPosFile

the output plain text file that stores the removed SNPs whose genomic positions are not in the imputation reference, ingoring SNP names.

snpSameNameDifPosFile

the output plain text file that stores the removed SNPs whose genomic positions are not in the imputation reference, taking SNP names into account.

reducedToSpecificfn

the prefix of PLINK files which are reduced in the number of SNPs and contain only SNPs prior to imputation.

specificDiffAllelefn

the prefix of PLINK files which added genotypes with different alleles other than the imputation to the file with the prefix: reducedToSpecificfn.

specificMissPosfn

the prefix of PLINK files which added genotypes with missing positions other than the imputation to the file with the prefix: specificDiffAllelefn.

specificDiffPosfn

the prefix of PLINK files which added genotypes with different positions other than the imputation to the file with the prefix: specificMissPosfn .

#### Value

Extracted genotypes from the imputed data which contain SNPs before imputation. On the basis of extracted genotypes, genotypes different from the imputation reference panel are appended, including SNPs with different alleles, missing positions, and different positions.

#### Author(s)

Junfang Chen

removedDoubleProbes

Remove duplicated SNPs

### **Description**

Remove duplicated SNPs that have same rs-names or duplicated genomic position.

### Usage

removedDoubleProbes(plink, inputPrefix, chipAnnoFile, chipType, outputSNPdupFile, outputPrefix) removedDoubleProbes 19

### **Arguments**

plink	an executable program in either the current working directory or somewhere in the command path.
inputPrefix	the prefix of the input PLINK binary files.
chipAnnoFile	an input chip annotation file. If the chip annotation file is not available for your study, it can be downloaded from http://www.well.ox.ac.uk/~wrayner/strand/.
chipType	a string name defines the type of the chip annotation file: 'SNPIDstudy', and 'rsIDstudy'. The detail is described in $prepareAnnoFile4affy$ .
outputSNPdupl	File a pure text file that stores the duplicated SNP IDs, which are detected by the use of the chip annotation file.
outputPrefix	the prefix of the output PLINK binary files.

### **Details**

Duplicated SNPs have two levels of meaning: 1.) SNPs have same rs-names but different versions of SNP ID in chip annotation file. e.g. SNP-A IDs for Affymetrix chip. 2.) SNPs with duplicated genomic position: the combination of base pair position and chromosomal location.

#### Value

The output PLINK binary files after removing duplicated SNP IDs.

# Author(s)

Junfang Chen

#### See Also

```
prepareAnnoFile4affy
```

```
## In the current working directory
bedFile <- system.file("extdata", "controlData.bed", package="Gimpute")
bimFile <- system.file("extdata", "controlData.bim", package="Gimpute")
famFile <- system.file("extdata", "controlData.fam", package="Gimpute")
chipAnnoFile <- system.file("extdata", "chipAnno.txt", package="Gimpute")
system(pasteO("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "controlData"
chipType <- "rsIDstudy"
outputSNPdupFile <- "snpDup.txt"
outputPrefix <- "removedDoubleProbes"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## removedDoubleProbes(plink, inputPrefix, chipAnnoFile,
## chipType, outputSNPdupFile, outputPrefix)</pre>
```

20 removedExclProbe

removedExclProbe Remove improper SNPs

### **Description**

Remove SNPs that may be duplicated, or with unexpected SNP names.

### Usage

```
removedExclProbe(plink, inputPrefix, excludedProbeIdsFile, outputPrefix)
```

#### **Arguments**

```
an executable program in either the current working directory or somewhere in the command path.

inputPrefix the prefix of the input PLINK binary files.

excludedProbeIdsFile

a pure text file that stores the SNP IDs, one per line, which need to be removed.

If it is null, then no SNPs are removed.

outputPrefix the prefix of the output PLINK binary files.
```

#### Details

excludedProbeIdsFile should be defined in a plain text file in advance. Improper SNPs such as AFFX, cnvi etc.. with unexpect format must be excluded.

#### Value

The output PLINK binary files after removing unwanted SNP IDs.

#### Author(s)

Junfang Chen

removedInstFhet 21

removedInstFhet

Remove subjects with abnormal autosomal heterozygosity deviation

### **Description**

Remove subjects with great autosomal heterozygosity deviation.

## Usage

```
removedInstFhet(plink, Fhet, inputPrefix, outputPrefix)
```

#### **Arguments**

plink an executable program in either the current working directory or somewhere in

the command path.

Fhet the cutoff of the autosomal heterozygosity deviation.

inputPrefix the prefix of the input PLINK binary files.
outputPrefix the prefix of the output PLINK binary files.

#### **Details**

If the cutoff of the autosomal heterozygosity deviation is set to be greater than 0.2, i.e.  $|Fhet| \ge 0.2$ , then this analysis will automatically skip haploid markers (male X and Y chromosome markers).

## Value

The output PLINK binary files after removing subjects with great autosomal heterozygosity deviation.

# Author(s)

Junfang Chen

```
## In the current working directory
bedFile <- system.file("extdata", "genoUpdatedData.bed", package="Gimpute")
bimFile <- system.file("extdata", "genoUpdatedData.bim", package="Gimpute")
famFile <- system.file("extdata", "genoUpdatedData.fam", package="Gimpute")
system(paste0("scp ", bedFile, bimFile, famFile, " ."))
Fhet <- 0.2
inputPrefix <- "genoUpdatedData"
outputPrefix <- "genoUpdatedData"
eutputPrefix <- "2_06_removedInstFhet"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## removedInstFhet(plink, Fhet, inputPrefix, outputPrefix)</pre>
```

22 removedInstMiss

removedInstMiss

Remove subjects with missing values

### Description

Remove Subjects or instances with missingness of greater than a certain threshold.

# Usage

```
removedInstMiss(plink, sampleMissCutOff, inputPrefix, outputPrefix)
```

# **Arguments**

```
plink an executable program in either the current working directory or somewhere in the command path.

sampleMissCutOff the cutoff of the missingness for removing subjects/instances.

inputPrefix the prefix of the input PLINK binary files.

outputPrefix the prefix of the output PLINK binary files.
```

### Value

The output PLINK binary files after removing subjects with a pre-defined removal cutoff.

#### Author(s)

Junfang Chen

```
## In the current working directory
bedFile <- system.file("extdata", "genoUpdatedData.bed", package="Gimpute")
bimFile <- system.file("extdata", "genoUpdatedData.bim", package="Gimpute")
famFile <- system.file("extdata", "genoUpdatedData.fam", package="Gimpute")
system(paste0("scp ", bedFile, bimFile, famFile, " ."))
sampleMissCutOff <- 0.02
inputPrefix <- "genoUpdatedData"
outputPrefix <- "genoUpdatedData"

**OutputPrefix <- "2_05_removedInstMiss"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## removedInstMiss(plink, sampleMissCutOff, inputPrefix, outputPrefix)</pre>
```

removedMaleHetX 23

removedMaleHetX

Remove male subjects with haploid heterozygous SNPs

### **Description**

Determine the frequency of male subjects that have heterozygous SNPs on chromosome X and a reasonable cutoff to remove those affect males, if chromosome X data exists.

### Usage

```
removedMaleHetX(plink, inputPrefix, hhSubjCutOff = 15, outputPrefix,
  outputSubjHetFile, outputRetainSubjectFile, outputHetSNPfile)
```

### Arguments

plink an executable program in either the current working directory or somewhere in

the command path.

inputPrefix the prefix of the input PLINK binary files.

hhSubjCutOff the cutoff for removing male subjects with haploid heterozygous SNPs on the

chromosome X. The default is 15.

outputPrefix the prefix of the output PLINK binary files.

outputSubjHetFile

the output pure text file that stores male subjects that have heterozygous SNPs with their frequency (if any), i.e. the number of .hh SNPs in this male. Lines are sorted by descending number.

outputRetainSubjectFile

the output pure text file that stores male subjects that have heterozygous SNPs with their frequency after subject removal (if any). Lines are sorted by descending number.

outputHetSNPfile

the output pure text file that stores all heterozygous SNPs with their frequency (the number of males for this SNP) , if any. Lines are sorted by descending number.

#### **Details**

A haploid heterozygous is a male genotype that is heterozygous, which could be an error given the haploid nature of the male X chromosome. In principle, one has to remove all males that have heterozygous SNPs on the chromosome X. However, too many males might be removed in some data sets. Therefore a small percentage of such males in the data set is allowed.

# Value

1.) The output PLINK binary files. 2.) A pure text file with two columns: heterozygous male subjects and their corresponding heterozygous SNPs. 3.) After subject removal, a pure text file consisting of two columns: heterozygous male subjects and their corresponding heterozygous SNPs. A pure text file with two columns: all heterozygous SNPs and their frequency.

### Author(s)

Junfang Chen

24 removedMendelErr

#### **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "genoUpdatedData.bed", package="Gimpute")
bimFile <- system.file("extdata", "genoUpdatedData.bim", package="Gimpute")</pre>
famFile <- system.file("extdata", "genoUpdatedData.fam", package="Gimpute")</pre>
system(paste0("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "genoUpdatedData"</pre>
hhSubjCutOff <- 15 ## can be tuned
outputPrefix <- "2_02_removedInstHetX"</pre>
outputSubjHetFile <- "2_02_instHetXfreqAll.txt"</pre>
outputRetainSubjectFile <- "2_02_instHetXfreqRetained.txt"</pre>
outputHetSNPfile <- "2_02_snpHHfreqAll.txt"</pre>
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## removedMaleHetX(plink, inputPrefix, hhSubjCutOff,
##
                      outputPrefix, outputSubjHetFile,
##
                      outputRetainSubjectFile, outputHetSNPfile)
```

removedMendelErr Check Mendel errors for family-based data

#### **Description**

Exclude subjects and/or genetic variants (SNPs) based on Mendel errors in the family data (trio/duo).

#### Usage

```
removedMendelErr(plink, inputPrefix, cutoffSubject = 0.05, cutoffSNP = 0.1,
  outputPrefix)
```

# Arguments

plink an executable program in either the current working directory or somewhere in

the command path.

inputPrefix the prefix of the input PLINK binary files.

cutoffSubject

the cutoff determines that families (subjects) with more than the predefined cutoff of Mendel errors by considering all SNPs will be removed. The default is

0.05.

cutoffSNP the cutoff indicates that SNPs with more than the predefined cutoff of Mendel

error rate will be excluded (i.e. based on the number of trios/duos). The default

is 0.1.

outputPrefix the prefix of the output PLINK binary files.

### Details

Do make sure that all your family relationships are correct in your input data before applying this function. The input PLINK data should have complete sex and group/outcome information. By default, trios and duos are both considered. If no family information is given at all (only founders), then this function will not remove any subjects or variants but give the warning showing that no duos or trios are present.

removedMonoSnp 25

#### Value

The output PLINK binary files.

#### Author(s)

Junfang Chen

#### **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "dataWithFamChr21.bed", package="Gimpute")
bimFile <- system.file("extdata", "dataWithFamChr21.bim", package="Gimpute")
famFile <- system.file("extdata", "dataWithFamChr21.fam", package="Gimpute")
system(paste0("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "dataWithFamChr21"
outputPrefix <- "removedMendelErr"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## removedMendelErr(plink, inputPrefix,
## cutoffSubject, cutoffSNP, outputPrefix)</pre>
```

removedMonoSnp

Exclude monomorphic SNPs

#### **Description**

Detect monomorphic SNPs from PLINK BIM file and exclude them if any.

# Usage

```
removedMonoSnp(plink, inputPrefix, outputPrefix, outputSNPfile)
```

### **Arguments**

plink an executable program in either the current working directory or somewhere in the command path.

inputPrefix the prefix of the input PLINK binary files.

 $\hbox{\tt outputPrefix the prefix of the output PLINK binary files after removing monomorphic SNPs.}\\$   $\hbox{\tt outputSNPfile}$ 

the output pure text file that stores the removed monomorphic SNPs, one per line, if any.

#### Value

The output PLINK binary files after removing monomorphic SNPs and a pure text file with removed monomorphic SNPs.

### Author(s)

Junfang Chen

26 removedParentIdsMiss

#### **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "alignedData.bed", package="Gimpute")
bimFile <- system.file("extdata", "alignedData.bim", package="Gimpute")
famFile <- system.file("extdata", "alignedData.fam", package="Gimpute")
system(paste0("scp ", bedFile, " ."))
system(paste0("scp ", bimFile, " ."))
system(paste0("scp ", famFile, " ."))
inputPrefix <- "alignedData"
outputPrefix <- "removedMonoSnp"
outputSNPfile <- "monoSNP.txt"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## removedMonoSnp(plink, inputPrefix, outputPrefix, outputSNPfile)</pre>
```

removedParentIdsMiss

Reset paternal and maternal codes

#### **Description**

Reset paternal and maternal codes of non-founders if parents not present. Replace the paternal ID and maternal ID of subjects (childs) by the value zero if the paternal ID and the maternal ID do not belong to any subject (parent) with the same family ID as the child.

### Usage

```
removedParentIdsMiss(plink, inputPrefix, outputPrefix)
```

# **Arguments**

```
plink an executable program in either the current working directory or somewhere in the command path.

inputPrefix the prefix of the input PLINK binary files.

outputPrefix the prefix of the output PLINK binary files.
```

#### **Details**

Do make sure that all your family relationships are correct in your input data before applying this function. By default, if parental IDs are provided for a sample, they are not treated as a founder even if neither parent is in the dataset. With no modifiers, –make-founders clears both parental IDs whenever at least one parent is not in the dataset, and the affected samples are now considered as founders.

### Value

The output PLINK binary files.

### Author(s)

Junfang Chen

### **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "genoUpdatedData.bed", package="Gimpute")
bimFile <- system.file("extdata", "genoUpdatedData.bim", package="Gimpute")
famFile <- system.file("extdata", "genoUpdatedData.fam", package="Gimpute")</pre>
system(paste0("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "genoUpdatedData"</pre>
outputPrefix <- "2_07_removedParentIdsMiss"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"</pre>
## removedParentIdsMiss(plink, inputPrefix, outputPrefix)
```

removedSnpFemaleChrXhweControl

Hardy Weinberg Equilibrium test for chromosome X SNPs in female controls.

## **Description**

Hardy Weinberg Equilibrium test for SNPs on the chromosome X in female controls.

#### Usage

```
removedSnpFemaleChrXhweControl(plink, inputPrefix, pval = 1e-06,
 outputPvalFile, outputSNPfile, outputPrefix)
```

### **Arguments**

an executable program in either the current working directory or somewhere in plink the command path. the prefix of the input PLINK binary files. inputPrefix the p-value cutoff for controlling HWE test in female control subjects. Only pval chromosome X SNPs are considered. The default value is 0.000001. outputPvalFile the output pure text file that stores chromosome X SNPs and their sorted HWE p-values.

outputSNPfile

the output pure text file that stores the removed SNPs, one per line.

outputPrefix the prefix of the output PLINK binary files.

### Value

The output PLINK binary files after HWE test on chromosomal X in female controls.

### Author(s)

Junfang Chen

### See Also

```
removedSnpHWEauto
```

#### **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "genoUpdatedData.bed", package="Gimpute")
bimFile <- system.file("extdata", "genoUpdatedData.bim", package="Gimpute")
famFile <- system.file("extdata", "genoUpdatedData.fam", package="Gimpute")
system(paste0("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "genoUpdatedData"
outputPvalFile <- "2_12_snpHwePvalfemaleXct.txt"
outputSNPfile <- "2_12_snpRemovedHweFemaleXct.txt"
outputPrefix <- "2_12_removedSnpHweFemaleX"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## removedSnpFemaleChrXhweControl(plink, inputPrefix, pval=0.000001,
outputPvalFile, outputSNPfile,
## outputPrefix)</pre>
```

removedSnpFemaleChrXmiss

remove chromosome X SNPs in females

### **Description**

Remove SNPs on the chromosome X with a pre-defined cutoff for missingness in females.

### Usage

```
removedSnpFemaleChrXmiss(plink, femaleChrXmissCutoff, inputPrefix, outputPrefix)
```

# **Arguments**

plink an executable program in either the current working directory or somewhere in the command path.

femaleChrXmissCutoff

the cutoff of the missingness in female chromosome X SNPs.

inputPrefix the prefix of the input PLINK binary files.

outputPrefix the prefix of the output PLINK binary files.

#### Value

The output PLINK binary files.

### Author(s)

Junfang Chen

removedSnpHetX 29

#### **Examples**

removedSnpHetX

Remove heterozygous SNPs in male chromosome X

### **Description**

Remove heterozygous SNPs in haploid male chromosome X only if chromosome X data exists.

# Usage

```
removedSnpHetX(plink, inputPrefix, hhCutOff, outputPrefix, outputHetSNPfile,
  outputRetainSNPfile)
```

### **Arguments**

plink an executable program in either the current working directory or somewhere in

the command path.

inputPrefix the prefix of the input PLINK binary files.

hhCutOff the cutoff for removing male haploid heterozygous SNPs on the chromosome

X. The default is 0.005.

outputPrefix the prefix of the output PLINK binary files.

outputHetSNPfile

the output pure text file that stores all heterozygous SNPs with their frequency (the number of males for the corresponding SNP), if any. Lines are sorted by descending number.

outputRetainSNPfile

the output pure text file that stores retained heterozygous SNPs with their frequency (the number of males for the corresponding SNP), if any. Lines are sorted by descending number.

#### **Details**

A haploid heterozygous is a male genotype that is heterozygous, which could be an error given the haploid nature of the male X chromosome. In principle, one has to remove all heterozygous SNPs of chromosome X in males. However, too many SNPs might be removed in some data sets. Therefore a small percentage of such SNPs in the data set is allowed.

#### Value

1.) The output PLINK binary files. 2.) A pure text files (if any) with two columns: SNPs and their corresponding frequency. 3.) After SNP removal, a pure text files (if any) with two columns: SNPs and their corresponding frequency.

### Author(s)

Junfang Chen

#### **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "genoUpdatedData.bed", package="Gimpute")
bimFile <- system.file("extdata", "genoUpdatedData.bim", package="Gimpute")
famFile <- system.file("extdata", "genoUpdatedData.fam", package="Gimpute")
system(pasteO("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "genoUpdatedData"
hhCutOff <- 0.005 ## can be tuned
outputPrefix <- "2_01_removedSnpHetX"
outputHetSNPfile <- "2_01_snpHHfreqAll.txt"
outputRetainSNPfile <- "2_01_snpHHfreqRetained.txt"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## removedSnpHetX(plink, inputPrefix, hhCutOff, outputPrefix,
outputHetSNPfile, outputRetainSNPfile)</pre>
```

removedSnpHWEauto Hardy Weinberg Equilibrium test for autosomal SNPs

### Description

Remove autosomal SNPs deviating from Hardy Weinberg Equilibrium (HWE).

outputPrefix the prefix of the output PLINK binary files.

### Usage

```
removedSnpHWEauto(groupLabel, plink, inputPrefix, pval = 1e-06,
  outputPvalFile, outputSNPfile, outputPrefix)
```

#### **Arguments**

```
a string value indicating the outcome label: "control", or, "case" or "caseControl" for both existing groups. For more details, see <a href="getGroupLabel">getGroupLabel</a>.

plink an executable program in either the current working directory or somewhere in the command path.

inputPrefix the prefix of the input PLINK binary files.

pval the p-value cutoff for controlling HWE test in either control or case subjects.

Only autosomal SNPs are considered. The default value is 0.000001.

outputPvalFile the output pure text file that stores autosomal SNPs and their sorted HWE p-values.

outputSNPfile the output pure text file that stores the removed SNPs, one per line.
```

removedSnpMiss 31

#### Value

The output PLINK binary files after HWE test on the autosome.

#### Author(s)

Junfang Chen

#### See Also

removedSnpFemaleChrXhweControl, getGroupLabel.

### **Examples**

removedSnpMiss

Remove SNPs with missing values

### **Description**

Remove SNPs with missingness of greater than a certain threshold.

# Usage

```
removedSnpMiss(plink, snpMissCutOff, inputPrefix, outputPrefix)
```

### **Arguments**

```
plink an executable program in either the current working directory or somewhere in the command path.

snpMissCutOff the cutoff of the missingness for removing SNPs.

inputPrefix the prefix of the input PLINK binary files.

outputPrefix the prefix of the output PLINK binary files.
```

#### Value

The output PLINK binary files after removing SNPs with pre-defined missing values.

#### Author(s)

Junfang Chen

#### **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "genoUpdatedData.bed", package="Gimpute")
bimFile <- system.file("extdata", "genoUpdatedData.bim", package="Gimpute")
famFile <- system.file("extdata", "genoUpdatedData.fam", package="Gimpute")
system(paste0("scp ", bedFile, bimFile, famFile, " ."))
snpMissCutOff <- 0.05
inputPrefix <- "genoUpdatedData"
outputPrefix <- "2_04_removedSnpMissPre"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## removedSnpMiss(plink, snpMissCutOff, inputPrefix, outputPrefix)</pre>
```

removedSnpMissDiff Remove SNPs with difference in SNP missingness between cases and controls.

#### **Description**

Remove SNPs with difference in SNP missingness between cases and controls. To test for differential call rates between cases and controls for each SNP

# Usage

```
removedSnpMissDiff(plink, inputPrefix, snpMissDifCutOff, outputPrefix,
  groupLabel)
```

# Arguments

```
plink an executable program in either the current working directory or somewhere in the command path.

inputPrefix the prefix of the input PLINK binary files.

snpMissDifCutOff the cutoff of the difference in missingness between cases and controls.

outputPrefix the prefix of the output PLINK binary files.

groupLabel a string value indicating the outcome label: "control", or, "case" or "caseControl" for both existing groups. For more details, see getGroupLabel.
```

### **Details**

Only if both case-control groups exist in the input genotype data, differential SNPs are removed.

#### Value

The output PLINK binary files.

#### Author(s)

Junfang Chen

#### See Also

```
getGroupLabel.
```

### **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "genoUpdatedData.bed", package="Gimpute")
bimFile <- system.file("extdata", "genoUpdatedData.bim", package="Gimpute")
famFile <- system.file("extdata", "genoUpdatedData.fam", package="Gimpute")
system(paste0("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "genoUpdatedData"
snpMissDifCutOff <- 0.02
outputPrefix <- "2_09_removedSnpMissDiff"
groupLabel <- "control"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## removedSnpMissDiff(plink, inputPrefix, snpMissDifCutOff,
outputPrefix, groupLabel)</pre>
```

removedSnpMissPostImp

Remove SNPs after post imputation

### **Description**

Remove SNPs which have a non missing value for less than a predefined number of instances.

### Usage

```
removedSnpMissPostImp(plink, inputPrefix, missCutoff, outRemovedSNPfile,
  outRetainSNPfile, outputPrefix)
```

#### **Arguments**

```
plink an executable program in either the current working directory or somewhere in the command path.

inputPrefix the prefix of the input PLINK binary files.

missCutoff the cutoff of the least number of instances for a SNP that is not missing. The default is 20.

outRemovedSNPfile the output file of SNPs with pre-defined missing values that are removed.

outRetainSNPfile the output file of SNPs that are retained.

outputPrefix the prefix of the PLINK binary files.
```

#### Value

The PLINK binary files after post imputation quality control and a pure text file contains SNPs with pre-defined missing values.

#### Author(s)

Junfang Chen

#### **Examples**

removedUnmapProbes Remove SNPs not in the chip annotation file

### **Description**

Check if all SNPs are included in the chip annotation file. If some of the input SNPs are not included, then remove them.

### Usage

```
removedUnmapProbes(plink, inputPrefix, chipAnnoFile, chipType, outputPrefix,
  outputSNPfile)
```

### **Arguments**

```
plink an executable program in either the current working directory or somewhere in the command path.

inputPrefix the prefix of the input PLINK binary files.

chipAnnoFile a pure text file that stores the chip annotation information.

chipType a string name defines the type of the chip annotation file: 'SNPIDstudy', and 'rsIDstudy'. The detail is described in prepareAnnoFile4affy.

outputPrefix the prefix of the output PLINK binary files.

outputSNPfile
```

a pure text file that stores the SNP IDs, one per line, which are not mapped to the chip annotation file.

#### **Details**

If the chip annotation file is not available for your study, you can download it from http://www.well.ox.ac.uk/~wrayner/str

#### Value

The output text file contains the removed SNP IDs, one per line. The PLINK binary files after removing unmapped SNP IDs.

#### Author(s)

Junfang Chen

#### See Also

```
prepareAnnoFile4affy
```

### **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "controlData.bed", package="Gimpute")
bimFile <- system.file("extdata", "controlData.bim", package="Gimpute")
famFile <- system.file("extdata", "controlData.fam", package="Gimpute")
chipAnnoFile <- system.file("extdata", "chipAnno.txt", package="Gimpute")
system(pasteO("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "controlData" ## Specify the input PLINK file prefix
outputSNPfile <- "1_07_removedUnmapProbes"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## removedUnmapProbes(plink, inputPrefix, chipAnnoFile, chipType,
## outputPrefix, outputSNPfile)</pre>
```

removedWrongAnceInst

Remove samples with incorrect ancestry

# Description

Remove samples with the incorrect ancestry or keep samples at your own chioce.

# Usage

```
removedWrongAnceInst(plink, inputPrefix, metaDataFile, ancestrySymbol,
  outputPrefix)
```

#### **Arguments**

```
plink an executable program in either the current working directory or somewhere in the command path.

inputPrefix the prefix of the input PLINK binary files.
```

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metaDataFile a pure text file that stores the meta information of the samples. This file must contain at least the following content (column names are in parentheses): family ID in the PLINK files (FID), individual ID in the PLINK files (IID), ID in the description files (descID), self identified ancestry (ance; e.g. AFR: African, AMR: Ad Mixed American, EAS: East Asian, EUR: European, SAS: South Asian), sex (sex; 1 = male, 2 = female), age (age), group (group; 0 = control/unaffected, 1 = case/affected). All unknown and missing values are represented by the value NA. Lines with a missing value for FID or IID are not contained.

ancestrySymbol

an indicator that shows the symbol of genetic ancestry. If it is null, then all samples are selected.

outputPrefix the prefix of the output PLINK binary files.

#### **Details**

ancestrySymbol, such as 'EUR' stands for the European, 'EAS' for East Asian. See the meta-DataFile for more details.

#### Value

The output PLINK binary files after checking the ancestry information.

#### Author(s)

Junfang Chen

### **Examples**

removedYMtSnp

Remove SNPs on the chromosome Y and mitochondrial DNA

### **Description**

Remove SNPs on the chromosome Y and mitochondrial DNA.

# Usage

```
removedYMtSnp(plink, inputPrefix, outputPrefix)
```

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### **Arguments**

```
plink an executable program in either the current working directory or somewhere in the command path.

inputPrefix the prefix of the input PLINK binary files.

outputPrefix the prefix of the output PLINK binary files.
```

#### **Details**

Note that if chromosome Y and mitochondrial DNA are available, they must be coded as 24 and 26, respectively.

#### Value

The output PLINK binary files after removing SNPs on the chromosome Y and mitochondrial DNA.

### Author(s)

Junfang Chen

### **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "controlData.bed", package="Gimpute")
bimFile <- system.file("extdata", "controlData.bim", package="Gimpute")
famFile <- system.file("extdata", "controlData.fam", package="Gimpute")
system(paste0("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "controlData" ## Specify the input PLINK file prefix
outputPrefix <- "1_11_removedYMtSnp"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## removedYMtSnp(plink, inputPrefix, outputPrefix)</pre>
```

removeNoGroupId

Remove samples without group information

### **Description**

Remove samples without group/outcome/phenotype information, which is coded as -9 in the PLINK .FAM file.

# Usage

```
removeNoGroupId(plink, inputPrefix, outputPrefix)
```

# **Arguments**

```
plink an executable program in either the current working directory or somewhere in the command path.

inputPrefix the prefix of the input PLINK binary files.

outputPrefix the prefix of the output PLINK binary files.
```

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#### Value

The output PLINK binary files after removing samples without group information.

### Author(s)

Junfang Chen

### **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "controlData.bed", package="Gimpute")
bimFile <- system.file("extdata", "controlData.bim", package="Gimpute")
famFile <- system.file("extdata", "controlData.fam", package="Gimpute")
system(pasteO("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "controlData" ## Specify the input PLINK file prefix
outputPrefix <- "1_04_removedNoGroupId"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## removeNoGroupId(plink, inputPrefix, outputPrefix)</pre>
```

removeOutlierByPCs Remove population outliers

# Description

Remove population outliers by using principle component analysis.

# Usage

```
removeOutlierByPCs(plink, gcta, inputPrefix, nThread = 20, cutoff = NULL,
  cutoffSign, inputPC4subjFile, outputPC4outlierFile, outputPCplotFile,
  outputPrefix)
```

### **Arguments**

plink	an executable program in either the current working directory or somewhere in the command path.
gcta	an executable program in either the current working directory or somewhere in the command path.
inputPrefix	the prefix of the input PLINK binary files.
nThread	the number of threads used for computation. The default is 20.
cutoff	the cutoff that distinguishes the outliers from ordinary population using PCA. If it is null, then there are no outliers or outliers are not required to be removed. The default is NULL.
cutoffSign	the cutoff sign: 'greater' or 'smaller' that determines if the outliers should be greater or smaller than the cutoff value.
inputPC4subjFile	
	the pure text file that stores all the subject IDs and their corresponding aircoval

the pure text file that stores all the subject IDs and their corresponding eigenvalues of the first two principle components.

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```
outputPC4outlierFile
```

the pure text file that stores the outlier IDs and their corresponding eigenvalues of the first two principle components.

```
outputPCplotFile
```

the plot file for visualizing the first two principle components of all subjects without population outliers.

outputPrefix the prefix of the output PLINK binary files.

#### **Details**

This function is used for removing population outliers. If the outliers are necessary to be removed, then one uses the eigenvalues from the first principle component as a criterion to find out the outliers by assigning an appropriate cutoff.

### Value

1.) The output PLINK binary files after outlier removal. 2.) The output pure text file (if any) for storing removed outlier IDs and their corresponding PCs. 3.) The plot file (if any) for visualizing the first two principle components after outlier removal.

### Author(s)

Junfang Chen

# **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "QCdata.bed", package="Gimpute")</pre>
bimFile <- system.file("extdata", "QCdata.bim", package="Gimpute")</pre>
famFile <- system.file("extdata", "QCdata.fam", package="Gimpute")</pre>
system(paste0("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "QCdata"
cutoff <- NULL ## no outlier to be removed
cutoffSign <- "greater" ## not used if cutoff == NULL</pre>
inputPC4subjFile <- "2_13_eigenvalAfterQC.txt"</pre>
outputPC4outlierFile <- "2_13_eigenval4outliers.txt"</pre>
outputPCplotFile <- "2_13_removedOutliers.png"</pre>
outputPrefix <- "2_13_removedOutliers"</pre>
## Not run: Requires an executable program PLINK and GCTA, e.g.
## plink <- "/home/tools/plink"</pre>
## gcta <- "/home/tools/gcta64"</pre>
## removeOutlierByPCs(plink, gcta, inputPrefix, nThread=20,
##
                        cutoff, cutoffSign, inputPC4subjFile,
##
                        outputPC4outlierFile, outputPCplotFile, outputPrefix)
```

removeSampID

Remove samples in PLINK files

# **Description**

Remove sample IDs that are useless such as duplicated or related IDs from PLINK binary files. These IDs are defined in a plain text file.

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### **Usage**

```
removeSampID(plink, removedSampIDFile, inputPrefix, outputPrefix)
```

### **Arguments**

plink an executable program in either the current working directory or somewhere in the command path.

removedSampIDFile a pure text file that stores the useless sample IDs, each ID per line. If it is null, then duplicate the input PLINK files as the output files.

inputPrefix the prefix of the input PLINK binary files.

outputPrefix the prefix of the output PLINK binary files.

#### Value

The output PLINK binary files after removing certain sample IDs.

### Author(s)

Junfang Chen

# **Examples**

renamePlinkBFile Rename PLINK binary files

# **Description**

Rename a set of PLINK binary files (.BED, .BIM and .FAM).

# Usage

```
renamePlinkBFile(inputPrefix, outputPrefix, action)
```

# Arguments

```
inputPrefix the prefix of the input PLINK binary files.
outputPrefix the prefix of the output PLINK binary files.
action a string indicating if the action is "copy" or "move".
```

### **Details**

The original input files can be retained using the action "copy" or removed by using "move".

# Value

Renamed PLINK binary files.

### Author(s)

Junfang Chen

# **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "controlData.bed", package="Gimpute")
bimFile <- system.file("extdata", "controlData.bim", package="Gimpute")
famFile <- system.file("extdata", "controlData.fam", package="Gimpute")
system(paste0("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "controlData"
outputPrefix <- "dataCtl"
outputPrefix <- "1_02_removedExclInst"
## renamePlinkBFile(inputPrefix, outputPrefix, action="move")</pre>
```

```
setHeteroHaploMissing
```

Set haploid heterozygous SNPs as missing

# **Description**

Set all heterozygous alleles of chromosome X SNPs in male as missing.

# Usage

```
setHeteroHaploMissing(plink, inputPrefix, outputPrefix)
```

# **Arguments**

```
plink an executable program in either the current working directory or somewhere in the command path.

inputPrefix the prefix of the input PLINK binary files.

outputPrefix the prefix of the output PLINK binary files.
```

# Value

The output PLINK binary files after setting haploid heterozygous SNPs as missing.

### Author(s)

Junfang Chen

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### **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "genoUpdatedData.bed", package="Gimpute")
bimFile <- system.file("extdata", "genoUpdatedData.bim", package="Gimpute")
famFile <- system.file("extdata", "genoUpdatedData.fam", package="Gimpute")
system(pasteO("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "genoUpdatedData"
outputPrefix <- "2_03_setHeteroHaploMissing"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## setHeteroHaploMissing(plink, inputPrefix, outputPrefix)</pre>
```

splitXchr

Split chromosome X into pseudoautosomal region and non-pseudoautosomal region.

# Description

Split chromosome X into pseudoautosomal region and non-pseudoautosomal region, if chromosome X data is available.

# Usage

```
splitXchr(plink, inputPrefix, outputPrefix)
```

# **Arguments**

plink an executable program in either the current working directory or somewhere in the command path.

inputPrefix the prefix of the input PLINK binary files.

outputPrefix the prefix of the output PLINK binary files.

# **Details**

Genomic coordinate system is on genome build hg19.

### Value

The output PLINK binary files after splitting chromosome X into pseudoautosomal region and non-pseudoautosomal region.

# Author(s)

Junfang Chen

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#### **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "controlData.bed", package="Gimpute")
bimFile <- system.file("extdata", "controlData.bim", package="Gimpute")
famFile <- system.file("extdata", "controlData.fam", package="Gimpute")
system(paste0("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "controlData" ## Specify the input PLINK file prefix
outputPrefix <- "1_10_splitXchr"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## removeSampID(plink, inputPrefix, outputPrefix)</pre>
```

updatedSnpInfo

Update the SNP information

### **Description**

Update SNP information including SNP name, base-pair position, chromosomal location and the strand information.

### Usage

```
updatedSnpInfo(plink, inputPrefix, chipAnnoFile, chipType, outputPrefix)
```

### **Arguments**

plink an executable program in either the current working directory or somewhere in the command path.

inputPrefix the prefix of the input PLINK binary files.

chipAnnoFile a pure text file that stores the chip annotation information. If the chip annotation file is not available for your study, it can be downloaded from http://www.well.ox.ac.uk/~wrayner/stra chipType a string name defines the type of the chip annotation file: 'SNPIDstudy', and 'rsIDstudy'. The detail is described in prepareAnnoFile4affy.

outputPrefix the prefix of the output PLINK binary files.

#### **Details**

The SNP information in the chip annotation file is used as the reference.

### Value

The output PLINK binary files after updating SNP information.

# Author(s)

Junfang Chen

### See Also

```
prepareAnnoFile4affy
```

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### **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "controlData.bed", package="Gimpute")
bimFile <- system.file("extdata", "controlData.bim", package="Gimpute")
famFile <- system.file("extdata", "controlData.fam", package="Gimpute")
chipAnnoFile <- system.file("extdata", "chipAnno.txt", package="Gimpute")
system(paste0("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "controlData"
chipType <- "rsIDstudy"
outputPrefix <- "updatedSnpInfo"
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## updatedSnpInfo(plink, inputPrefix, chipAnnoFile,
## chipType, outputPrefix, outputSNPfile)</pre>
```

updateGenoInfo

Update genotype information

# **Description**

Update genotype information of the original PLINK binary files involving subject metadata information remapping and SNP information rearrangement and conversion according to the annotation file.

# Usage

```
updateGenoInfo(plink, inputPrefix, metaDataFile, removedSampIDFile,
  ancestrySymbol, excludedProbeIdsFile, chipAnnoFile, chipType, outputPrefix,
  keepInterFile = TRUE)
```

# **Arguments**

plink

an executable program in either the current working directory or somewhere in the command path.

inputPrefix the prefix of the input PLINK binary files.

-

metaDataFile a pure text file that stores the meta information of the samples. This file must contain at least the following content (column names are in parentheses): family ID in the PLINK files (FID), individual ID in the PLINK files (IID), ID in the description files (descID), self identified ancestry (ance; e.g. AFR: African, AMR: Ad Mixed American, EAS: East Asian, EUR: European, SAS: South Asian), sex (sex; 1 = male, 2 = female), age (age), group (group; 0 = control/unaffected, 1 = case/affected). All unknown and missing values are represented by the value NA. Lines with a missing value for FID or IID are not contained.

removedSampIDFile

a pure text file that stores the useless sample IDs, each ID per line. If it is null, then duplicate the input PLINK files from the last step as the output files.

ancestrySymbol

an indicator that shows the symbol of genetic ancestry. If it is null, then all samples are selected.

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```
excludedProbeIdsFile

a pure text file that stores the SNP IDs, one per line, which need to be removed.

If it is null, no SNPs are removed.

chipAnnoFile a pure text file that stores the chip annotation information.

chipType a string name defines the type of the chip annotation file: 'SNPIDstudy', and 'rsIDstudy'. The detail is described in prepareAnnoFile4affy.
```

 $\verb"outputPrefix" the prefix of the output PLINK binary files.$ 

keepInterFile

a logical value indicating if the intermediate processed files should be kept or not. The default is TRUE.

#### **Details**

The original PLINK files are implicitly processed by the following steps: 1.) remove duplicated subjects; 2.) update group ID and sex information; 3.) remove not labelled subjects; 4.) remove subjects with wrong ancestry; 5.) remove incorrectly annotated SNPs; 6.) remove SNPs that are not in the annotation file; 7.) remove duplicated SNPs; 8.) update SNP genomic position and strand information; 9.) split chromosome X into pseudoautosomal region (PAR) and non-PAR; 10.) remove SNPs on the chromosome Y and mitochondrial DNA. The metadata information file and the chip annotation file are used as the reference for the update. If the chip annotation file is not available for your study, it can be downloaded from http://www.well.ox.ac.uk/~wrayner/strand/.

#### Value

The output PLINK binary files after genotype information remapping.

# Author(s)

Junfang Chen

### References

Purcell, Shaun, et al. PLINK: a tool set for whole-genome association and population-based linkage analyses. The American Journal of Human Genetics 81.3 (2007): 559-575.

### **Examples**

```
## In the current working directory
bedFile <- system.file("extdata", "controlData.bed", package="Gimpute")</pre>
bimFile <- system.file("extdata", "controlData.bim", package="Gimpute")</pre>
famFile <- system.file("extdata", "controlData.fam", package="Gimpute")</pre>
system(paste0("scp ", bedFile, bimFile, famFile, " ."))
inputPrefix <- "controlData"</pre>
metaDataFile <- system.file("extdata", "1_01_metaData.txt",</pre>
                              package="Gimpute")
excludedProbeIdsFile <- system.file("extdata", "excludedProbeIDs.txt",</pre>
                                       package="Gimpute")
removedSampIDFile <- system.file("extdata", "excludedSampIDsV1.txt",
                                    package="Gimpute")
chipAnnoFile <- system.file("extdata", "chipAnno.txt", package="Gimpute")</pre>
ancestrySymbol <- "EUR"</pre>
outputPrefix <- "1_11_removedYMtSnp"</pre>
metaDataFile <- "1_01_metaData.txt"</pre>
chipType <- "rsIDstudy"</pre>
```

```
## Not run: Requires an executable program PLINK, e.g.
## plink <- "/home/tools/plink"
## updateGenoInfo(plink, inputPrefix, metaDataFile, removedSampIDFile,
## ancestrySymbol, excludedProbeIdsFile, chipAnnoFile,
## chipType, outputPrefix, keepInterFile=TRUE)</pre>
```

updateGroupIdAndSex

Update group and geneder information

### **Description**

Replace group and gender information in the PLINK binary files by using the information from the metadata file.

#### Usage

```
updateGroupIdAndSex(plink, inputPrefix, metaDataFile, outputPrefix)
```

### **Arguments**

plink an executable program in either the current working directory or somewhere in

the command path.

inputPrefix the prefix of the input PLINK binary files.

metaDataFile a pure text file that stores the meta information of the samples. This file must contain at least the following content (column names are in parentheses): family

ID in the PLINK files (FID), individual ID in the PLINK files (IID), ID in the description files (descID), self identified ancestry (ance; e.g. AFR: African, AMR: Ad Mixed American, EAS: East Asian, EUR: European, SAS: South Asian), sex (sex; 1 = male, 2 = female), age (age), group (group; 0 = control/unaffected, 1 = case/affected). All unknown and missing values are represented by the value

NA. Lines with a missing value for FID or IID are not contained.

outputPrefix the prefix of the output PLINK binary files.

### **Details**

Find the shared sample IDs between PLINK input files and metadata file. Use the information from the metadata file as the reference and update the group information/the outcome in the PLINK file. Group label should be 1 and 2. (1=unaff, 2=aff, 0=miss); missing phenotype will be indicated as -9.

### Value

The output PLINK binary files after updating the gender and grouping information.

### Author(s)

Junfang Chen

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