

# Package ‘simGWAS’

August 22, 2019

**Type** Package

**Title** Fast Simulation of Large Case-Control GWAS Summary Statistics

**Version** 0.2.0-2

**Date** 2019-08-19

**Author** Mary Fortune [aut],  
Chris Wallace [aut, cre],  
Marcus Klarqvist [ctb]

**Maintainer** Chris Wallace <cew54@cam.ac.uk>

**Description** Simulating output from a case-control genome wide association study (GWAS) with a given causal model. Fortune and Wallace (2019) <doi:10.1093/bioinformatics/bty898>.

**License** GPL

**LazyLoad** yes

**OS\_type** unix

**Collate** 'simGWAS.R' 'compute\_gamma0.R' 'vbeta.R' 'zscore.R'  
'extractsnps.R' 'make\_dataset.R' 'make\_GenoProbList.R'  
'RcppExports.R'

**LinkingTo** Rcpp

**Imports** Rcpp, dplyr, combinat, corpcor, mvtnorm

**RoxygenNote** 6.1.1

**Suggests** knitr, rmarkdown, testthat

**VignetteBuilder** knitr

**URL** <https://github.com/chr1swallace/simGWAS>

**BugReports** <https://github.com/chr1swallace/simGWAS/issues>

**NeedsCompilation** yes

**Repository** CRAN

**Date/Publication** 2019-08-22 08:20:02 UTC

**R topics documented:**

compute_gamma0 . . . . .	2
compute_gamma0_PW . . . . .	3
est_statistic . . . . .	3
est_zscore . . . . .	4
expected_vbeta . . . . .	5
expected_z_score . . . . .	6
fake_freq . . . . .	7
make_GenoProbList . . . . .	8
simulated_vbeta . . . . .	8
simulated_z_null . . . . .	9
simulated_z_score . . . . .	10
which_genotypes . . . . .	11

<b>Index</b>	<b>12</b>
--------------	-----------

---

compute_gamma0	<i>Compute gamma0, given haplotype frequencies</i>
----------------	--

---

**Description**

Compute the value of gamma\_0 from gamma1,...,gamma\_m

**Usage**

```
compute_gamma0(N0, N1, W, gamma.W, freq)
```

**Arguments**

N0	number of samples with Y=0
N1	number of samples with Y=1
W	The causal SNPs
gamma.W	The odds ratios of effect for each CV
freq	Frequencies of SNP appearances (computed using snphap)

**Details**

Note: assume we must compute the distribution of W in the controls

**Value**

The value of gamma0

**Author(s)**

Mary Fortune

---

compute\_gamma0\_PW      *Compute gamma0 given distribution of W*

---

**Description**

Compute the value of gamma\_0 from gamma1,...,gamma\_m

**Usage**

compute\_gamma0\_PW(N0, N1, W, gamma.W, PWgY0)

**Arguments**

N0	number of samples with Y=0
N1	number of samples with Y=1
W	The causal SNPs
gamma.W	The odds ratios of effect for each CV
PWgY0	the distribution of causal snps in controls

**Details**

Note: assume we know the distribution of W in the controls

**Value**

The value of gamma0

**Author(s)**

Mary Fortune

---

est\_statistic      *estimate Z score at a single SNP*

---

**Description**

Wrapper function to run est\_zscore for all snps in snps

**Usage**

est\_statistic(N0, N1, snps, W, gamma.W, freq, GenoProbList)

**Arguments**

N0	The number of Y=0
N1	The number of Y=1
snps	The snps at which we wish to compute the expected Z Score
W	The true causal SNPs (these need not be in "snps")
gamma.W	The odds ratios of effect of the true causal SNPs (not including gamma0, the intercept term)
freq	Frequencies of SNP appearances (computed using snphap)
GenoProbList	An list of objects giving the probability of seeing each X,W genotype vector

**Details**

Assumes we have a list, GenoProbList, giving the GenoProb values for each X.

**Value**

The expected Z Score for SNP X, assuming the causal SNPs are W

**Author(s)**

Mary Fortune and Chris Wallace

---

est_zscore	<i>estimate Z score at a single SNP</i>
------------	---

---

**Description**

Estimates the expected Z Score for a single SNP

**Usage**

```
est_zscore(N0, N1, Ufactor, powerfactor, freq, GenoProbXW)
```

**Arguments**

N0	The number of Y=0
N1	The number of Y=1
Ufactor	The constant factor used to compute the expectation of U
powerfactor	The constant factor used to compute the expectation of the genotype of X to some power
freq	Frequencies of SNP appearances (computed using snphap)
GenoProbXW	An object giving the probability of seeing each X,W genotype vector

**Details**

Assumes the input CVs, and the relationship, gamma, between them and the trait of interest  
 Assumes we have already generated GenoProbXW for all X

**Value**

The expected Z Score for SNP X, assuming the causal SNPs are W

**Author(s)**

Mary Fortune and Chris Wallace

---

expected_vbeta	<i>Estimate the expected variance of beta</i>
----------------	---

---

**Description**

Estimate the expected variance of beta. This is approximately  $\text{expected}(1/\text{var}(U))$ .

**Usage**

```
expected_vbeta(N0, N1, snps, W, gamma.W, freq,
  GenoProbList = make_GenoProbList(snps = snps, W = W, freq = freq))
```

**Arguments**

- N0                    The number of Y=0
- N1                    The number of Y=1
- snps                   The snps at which we wish to compute the expected Z Score
- W                      The true causal SNPs (these need not be in "snps")
- gamma.W               The log odds ratios of effect of the true causal SNPs (not including gamma0, the intercept term)
- freq                   Haplotype frequencies as a data.frame, with column Probability indicating relative frequency in controls.
- GenoProbList         An list of objects giving the probability of seeing each X,W genotype vector. This can be calculated within the function if no value supplied, or you can pass a pre-calculated version

**Details**

Assumes we have a list, GenoProbList, giving the GenoProb values for each X.

**Value**

The expected variance of beta for each SNP X, assuming the causal SNPs are W

**Author(s)**

Mary Fortune and Chris Wallace

**Examples**

```
freq=fake_freq(nhaps=100,nsnps=5) # fake haplotype frequency data
EVB=expected_vbeta(N0=1000,N1=2000,snps=paste0("s",1:5),
                  W="s1",gamma.W=log(1.5),freq=freq)
EVB # causal variant is SNP 1, with OR 1.5
```

---

expected_z_score	<i>Compute vector of expected Z Scores</i>
------------------	--

---

**Description**

Compute vector of expected Z scores

**Usage**

```
expected_z_score(N0, N1, snps, W, gamma.W, freq,
                 GenoProbList = make_GenoProbList(snps = snps, W = W, freq = freq))
```

**Arguments**

N0	The number of Y=0
N1	The number of Y=1
snps	The snps at which we wish to compute the expected Z Score
W	The true causal SNPs (these need not be in "snps")
gamma.W	The log odds ratios of effect of the true causal SNPs (not including gamma0, the intercept term)
freq	Haplotype frequencies as a data.frame, with column Probability indicating relative frequency in controls.
GenoProbList	An list of objects giving the probability of seeing each X,W genotype vector. This can be calculated within the function if no value supplied, or you can pass a pre-calculated version

**Value**

The expected Z Score for all snps in snps, assuming the causal SNPs are W

**Author(s)**

Mary Fortune and Chris Wallace

**Examples**

```
freq=fake_freq(nhaps=100,nsnps=5) # fake haplotype frequency data
EZ=expected_z_score(N0=1000,N1=2000,snps=paste0("s",1:5),
                    W="s1",gamma.W=log(1.5),freq=freq)
EZ # causal variant is SNP 1, with OR 1.5
```

---

fake_freq	<i>fake haplotype frequencies</i>
-----------	-----------------------------------

---

**Description**

create a fake haplotype frequency dataset

**Usage**

```
fake_freq(nhaps = 100, nsnps = 10)
```

**Arguments**

nhaps	number of haplotypes
nsnps	number of snps

**Details**

no attempt is made at biological realism, this is purely for testing code

**Value**

data.frame of 1,2, nhaps x nsamples + frequency column

**Author(s)**

Chris Wallace

**Examples**

```
freq=fake_freq(nhaps=100,nsnps=5)
dim(freq)
head(freq)
```

---

make\_GenoProbList      *make\_GenoProbList*

---

### Description

compute a list, GenoProbList, giving the GenoProb values for each X.

### Usage

```
make_GenoProbList(snps, W, freq)
```

### Arguments

snps	Index of SNPs at which we wish to compute the expected Z Score
W	Index of true causal SNPs (these need not be in "snps")
freq	Frequencies of SNP appearances (computed using snphap)

### Value

The the GenoProb values for each X

### Author(s)

Mary Fortune and Chris Wallace

### Examples

```
freq=fake_freq() # fake haplotype frequency data
problast=make_GenoProbList(1:2,W=1,freq)
```

---

simulated\_vbeta      *Compute a simulated var(beta)*

---

### Description

Simulate var(beta)

### Usage

```
simulated_vbeta(N0, N1, snps, W, gamma.W, freq,
  GenoProbList = make_GenoProbList(snps = snps, W = W, freq = freq),
  nrep = 1)
```



**Arguments**

N0	The number of Y=0
N1	The number of Y=1
snps	The snps at which we wish to compute the expected Z Score
W	The true causal SNPs (these need not be in "snps")
gamma.W	The log odds ratios of effect of the true causal SNPs (not including gamma0, the intercept term)
freq	Haplotype frequencies as a data.frame, with column Probability indicating relative frequency in controls.
GenoProbList	An list of objects giving the probability of seeing each X,W genotype vector. This can be calculated within the function if no value supplied, or you can pass a pre-calculated version
nrep	Number of replicates (simulated vectors of Z scores) under this scenario. Default=1

**Details**

Assumes we have a list, GenoProbList, giving the GenoProb values for each X.

**Value**

A simulated variance of beta for each SNP X, assuming the causal SNPs are W

**Author(s)**

Mary Fortune and Chris Wallace

**Examples**

```
freq=fake_freq(nhaps=100,nsnps=5) # fake haplotype frequency data
VB=simulated_vbeta(N0=1000,N1=2000,snps=paste0("s",1:5),
                  W="s1",gamma.W=log(1.5),freq=freq)
VB # causal variant is SNP 1, with OR 1.5
```

---

simulated\_z\_null      *Compute a NULL simulated Z Score*

---

**Description**

Compute matrix of simulated Z scores about expected values of 0 - ie under a null of no association at any SNP

**Usage**

```
simulated_z_null(snps, freq, nrep = 1)
```

**Arguments**

snps	The snps at which we wish to compute the expected Z Score
freq	Haplotype frequencies as a data.frame, with column Probability indicating relative frequency in controls.
nrep	Number of replicates (simulated vectors of Z scores) under this scenario. Default=1

**Author(s)**

Mary Fortune and Chris Wallace

**Examples**

```
freq=fake_freq(nhaps=100,nsnps=5) # fake haplotype frequency data
Z=simulated_z_null(snps=paste0("s",1:5),freq=freq,nrep=3)
Z # no causal variants
```

---

simulated_z_score	<i>Compute a simulated Z Score</i>
-------------------	------------------------------------

---

**Description**

Compute matrix of simulated Z scores

**Usage**

```
simulated_z_score(N0, N1, snps, W, gamma.W, freq,
  GenoProbList = make_GenoProbList(snps = snps, W = W, freq = freq),
  nrep = 1)
```

**Arguments**

N0	The number of Y=0
N1	The number of Y=1
snps	The snps at which we wish to compute the expected Z Score
W	The true causal SNPs (these need not be in "snps")
gamma.W	The log odds ratios of effect of the true causal SNPs (not including gamma0, the intercept term)
freq	Haplotype frequencies as a data.frame, with column Probability indicating relative frequency in controls.
GenoProbList	An list of objects giving the probability of seeing each X,W genotype vector. This can be calculated within the function if no value supplied, or you can pass a pre-calculated version
nrep	Number of replicates (simulated vectors of Z scores) under this scenario. Default=1

**Author(s)**

Mary Fortune and Chris Wallace

**Examples**

```
freq=fake_freq(nhaps=100,nsnps=5) # fake haplotype frequency data
Z=simulated_z_score(N0=1000,N1=2000,snps=paste0("s",1:5),
                    W="s1",gamma.W=log(1.5),freq=freq,nrep=3)
Z # causal variant is SNP 1, with OR 1.5
```

---

*which\_genotypes*      *maps haplotype pairs to genotypes*

---

**Description**

computes a matrix telling us which haplotype pairs correspond to which genotypes

**Usage**

```
which_genotypes(nsnps)
```

**Arguments**

nsnps            The number of SNPs

**Details**

Internal function

**Author(s)**

Mary Fortune

# Index

`compute_gamma0`, 2  
`compute_gamma0_PW`, 3

`est_statistic`, 3  
`est_zscore`, 4  
`expected_vbeta`, 5  
`expected_z_score`, 6

`fake_freq`, 7

`make_GenoProbList`, 8

`simulated_vbeta`, 8  
`simulated_z_null`, 9  
`simulated_z_score`, 10

`which_genotypes`, 11