

Tutorial: Using *GeneticsBase*

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```
> options(width = 90)
```

1 Introduction

This vignette was created as a tutorial for the 2007 BioConductor User's Conference held in Seattle, WA, USA during August 2007, and was presented by Dr. Warnes and Dr. Lazarus. The material is structured as a tutorial with a small example data set (8184 Markers x 180 Subjects belonging to 50 Families) .

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3 Preliminaries

3.1 Install RGenetics packages and dependencies

For MS-Windows it is necessary to manually install dependencies from CRAN

```
> install.packages(c("xtable", "combinat", "gdata", "gplots", "mvtnorm"),  
+   dep = TRUE)
```

Now to install the necessary packages:

```
> repos <- c("http://www.warnes.net/bioc2007/", "http://cran.fhcrc.org")  
> install.packages(c("GeneticsBase", "GeneticsDesign", "fbat"), repos = repos,  
+   type = "source", dep = TRUE)
```

3.2 Load the libraries

```
> library(GeneticsBase)  
> library(GeneticsDesign)  
> library(fbat)
```

4 Loading Data

4.1 Read data from files

Load the full data:

```
> hm.a <- readGenes(gfile = "hmCEU_YRI_chr22_ALLfbat.ped", gformat = "fbat")
```

```
Reading 8184 markers and 180 subjects from `hmCEU_YRI_chr22_ALLfbat.ped' ...  
generating 'geneSet' object...
```

```
100 200 300 400 500 600 700 800 900 1000  
1100 1200 1300 1400 1500 1600 1700 1800 1900 2000  
2100 2200 2300 2400 2500 2600 2700 2800 2900 3000  
3100 3200 3300 3400 3500 3600 3700 3800 3900 4000  
4100 4200 4300 4400 4500 4600 4700 4800 4900 5000  
5100 5200 5300 5400 5500 5600 5700 5800 5900 6000  
6100 6200 6300 6400 6500 6600 6700 6800 6900 7000  
7100 7200 7300 7400 7500 7600 7700 7800 7900 8000  
8100 Successfully read the pedigree file `hmCEU_YRI_chr22_ALLfbat.ped' .
```

```
Number of Markers: 8184  
Number of Subjects: 180  
Number of Families: 50
```

```
> print(hm.a)
```

```
geneSet object  
-----
```

```
Number of Markers:      8184  
Number of Observations: 180
```

Sample variables: family, pid, father, mother, sex, affected

Genetic data:

| | 1334.1 | 1334.10 | 1334.11 | 1334.12 | 1334.13 | 1334.2 | 74.3 | 77.1 | 77.2 | |
|-----------------------|--------|---------|---------|---------|---------|--------|------|------|------|-----|
| 22_14884399_rs4911642 | 4/4 | 4/4 | 4/4 | 4/4 | 4/4 | 4/4 | ... | 4/4 | 4/4 | 2/4 |
| 22_15298335_rs2027653 | 2/2 | 2/4 | 2/4 | 4/4 | 2/4 | 4/4 | ... | 4/4 | 2/4 | 2/2 |
| 22_15412698_rs5747620 | 2/2 | 2/2 | 2/2 | 2/4 | 2/4 | 4/4 | ... | 2/4 | 2/4 | 2/2 |
| 22_15434720_rs9605903 | 4/4 | 4/4 | 4/4 | 2/4 | 4/4 | 2/4 | ... | 4/4 | 4/4 | 4/4 |
| 22_15447504_rs5747968 | 3/4 | 4/4 | 3/4 | 3/4 | 4/4 | 3/4 | ... | 4/4 | 4/4 | 4/4 |
| 22_15452483_rs2236639 | 3/3 | 3/3 | 3/3 | 3/3 | 3/3 | 3/3 | ... | 3/3 | 3/3 | 3/3 |
| | . | . | . | . | . | . | . | . | . | . |
| | . | . | . | . | . | . | . | . | . | . |
| | . | . | . | . | . | . | . | . | . | . |
| 22_49497339_rs5770820 | 1/3 | <NA> | 1/1 | 3/3 | 3/3 | 3/3 | ... | 3/3 | 3/3 | 3/3 |
| 22_49498590_rs6010061 | 2/4 | 2/2 | 4/4 | 2/2 | 2/2 | 2/2 | ... | 2/4 | 2/4 | 4/4 |
| 22_49510004_rs715586 | 2/2 | 2/2 | 2/2 | 2/2 | 2/2 | 2/2 | ... | 2/2 | 2/2 | 2/2 |
| 22_49512530_rs8137951 | 1/3 | 1/3 | 1/1 | 3/3 | 3/3 | 3/3 | ... | 3/3 | 1/3 | 1/3 |
| 22_49518559_rs756638 | 3/3 | 3/3 | 3/3 | 1/3 | 3/3 | 3/3 | ... | 1/3 | 3/3 | 1/3 |
| 22_49522492_rs3810648 | 1/1 | 1/1 | 1/1 | 1/1 | 1/1 | 1/1 | ... | 1/3 | 1/1 | 1/1 |
| | 77.3 | 9.1 | 9.2 | | | | | | | |
| 22_14884399_rs4911642 | 2/4 | <NA> | 2/4 | | | | | | | |
| 22_15298335_rs2027653 | 4/4 | 4/4 | 4/4 | | | | | | | |
| 22_15412698_rs5747620 | 4/4 | 2/4 | 4/4 | | | | | | | |
| 22_15434720_rs9605903 | 4/4 | 4/4 | 4/4 | | | | | | | |
| 22_15447504_rs5747968 | 4/4 | 4/4 | 4/4 | | | | | | | |
| 22_15452483_rs2236639 | 3/3 | 3/3 | 3/3 | | | | | | | |
| | . | . | . | | | | | | | |
| | . | . | . | | | | | | | |
| | . | . | . | | | | | | | |
| 22_49497339_rs5770820 | 3/3 | 3/3 | 3/3 | | | | | | | |
| 22_49498590_rs6010061 | 2/4 | 4/4 | 4/4 | | | | | | | |
| 22_49510004_rs715586 | 2/2 | 2/2 | 2/2 | | | | | | | |
| 22_49512530_rs8137951 | 1/3 | 1/1 | 1/3 | | | | | | | |
| 22_49518559_rs756638 | 3/3 | 1/3 | 1/3 | | | | | | | |
| 22_49522492_rs3810648 | 1/1 | 1/3 | 1/1 | | | | | | | |

We'll also need just the founders later:

```
> hm.f <- readGenes(gfile = "hmCEU_YRI_chr22_Foundersfbat.ped", gformat = "fbat")
```

```
Reading 8184 markers and 120 subjects from `hmCEU_YRI_chr22_Foundersfbat.ped` ...  
generating 'geneSet' object...
```

```
100 200 300 400 500 600 700 800 900 1000  
1100 1200 1300 1400 1500 1600 1700 1800 1900 2000  
2100 2200 2300 2400 2500 2600 2700 2800 2900 3000  
3100 3200 3300 3400 3500 3600 3700 3800 3900 4000  
4100 4200 4300 4400 4500 4600 4700 4800 4900 5000  
5100 5200 5300 5400 5500 5600 5700 5800 5900 6000  
6100 6200 6300 6400 6500 6600 6700 6800 6900 7000  
7100 7200 7300 7400 7500 7600 7700 7800 7900 8000  
8100 Successfully read the pedigree file `hmCEU_YRI_chr22_Foundersfbat.ped`.
```

```
Number of Markers: 8184
Number of Subjects: 120
Number of Families: 50
```

For the purpose of speeding execution of examples, we'll also create a smaller subset of 100 markers from the original file.

```
> hm.a2 <- hm.a[1:100, ]
```

4.2 Error check the loaded data

Count frequencies of missing genotypes (requires 26 seconds on my MacBook Pro)

Number of missing genotypes per subject:

```
> mG <- missGFreq(hm.a2, founderOnly = TRUE, quiet = FALSE)
```

converting geneSet object to numerical matrix...

counting frequencies of missing genotypes...

```
> head(mG$nMissSubjects)
```

```
      00 0* *0
subject2 3 0 0
subject3 2 0 0
subject4 4 0 0
subject5 0 0 0
subject8 1 0 0
subject9 3 0 0
```

Column headers:

00 missing both alleles

0* 1st allele missing while the 2nd allele is not missing

*0 1st allele is not missing while the 2nd allele is missing

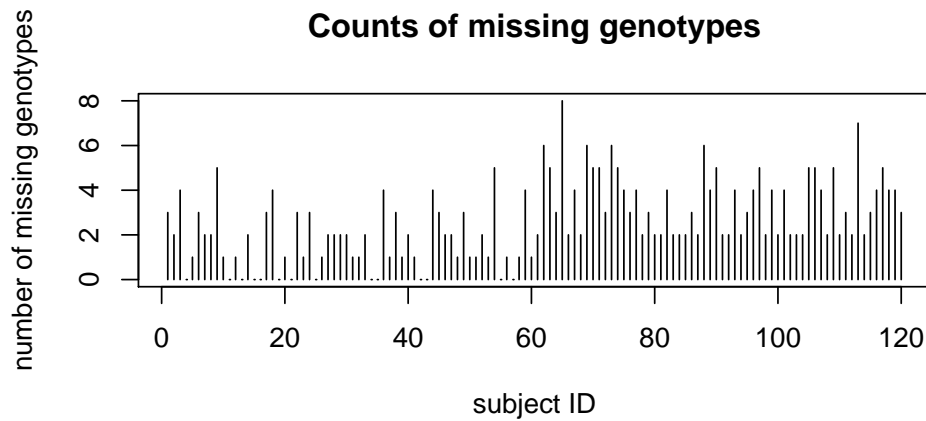
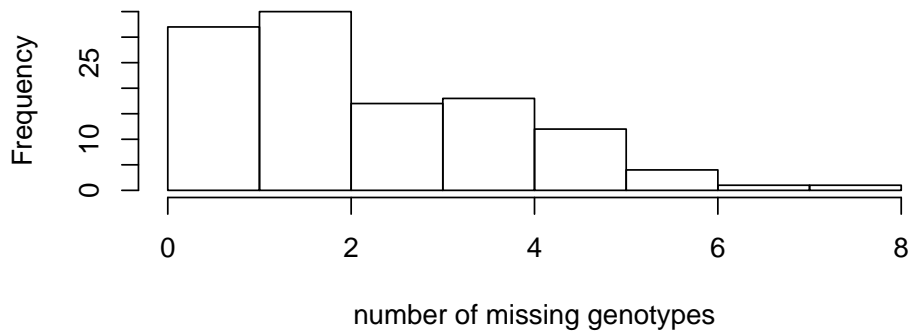
Number of missing genotypes per marker:

```
> head(mG$nMissMarkers)
```

```
      00 0* *0
22_14884399_rs4911642 13 0 0
22_15298335_rs2027653  1 0 0
22_15412698_rs5747620  0 0 0
22_15434720_rs9605903  0 0 0
22_15447504_rs5747968  0 0 0
22_15452483_rs2236639  0 0 0
```

Plot counts of missing genotypes:

```
> par(mfrow = c(2, 1))
> hist(mG$nMissSubjects[, 1], main = "", xlab = "number of missing genotypes")
> plot(1:nrow(mG$nMissSubjects), mG$nMissSubjects[, 1], xlab = "subject ID",
+      ylab = "number of missing genotypes", type = "h")
> title("Counts of missing genotypes")
> par(mfrow = c(1, 1))
```



5 Descriptive Statistics

1. Summary of allele/genotype frequency
2. HWE test
3. Visualize Disequilibrium

Basic data quality checks for markers. Column headings are:

ObsHET observed proportion of heterozygous genotypes per marker

PredHET predicted proportion of heterozygous genotypes per marker

HWpval pvalues of Hardy-Weinberg test per marker

pGeno percentage of non-missing genotypes for markers

MAF minor allele frequencies. missing alleles are excluded from calculation

Rating 1 if passes HW test; 0 if failed HW test.

```
> cM <- checkMarkers(hm.a2)
> head(cM)
```

| | Name | Position | ObsHET | PredHET | Hwpval |
|-----------------------|-----------------------|-----------|--------------|-----------|------------|
| 22_14884399_rs4911642 | 22_14884399_rs4911642 | | ? 0.30841121 | 0.2608525 | 0.05930340 |
| 22_15298335_rs2027653 | 22_15298335_rs2027653 | | ? 0.34453782 | 0.3892734 | 0.20997430 |
| 22_15412698_rs5747620 | 22_15412698_rs5747620 | | ? 0.50000000 | 0.4994444 | 0.99027790 |
| 22_15434720_rs9605903 | 22_15434720_rs9605903 | | ? 0.16666667 | 0.2061111 | 0.03604633 |
| 22_15447504_rs5747968 | 22_15447504_rs5747968 | | ? 0.23333333 | 0.2665278 | 0.17246954 |
| 22_15452483_rs2236639 | 22_15452483_rs2236639 | | ? 0.09166667 | 0.1171875 | 0.01704962 |
| | pGeno | MAF | Rating | | |
| 22_14884399_rs4911642 | 89.16667 | 0.1542056 | 1 | | |
| 22_15298335_rs2027653 | 99.16667 | 0.2647059 | 1 | | |
| 22_15412698_rs5747620 | 100.00000 | 0.4833333 | 1 | | |
| 22_15434720_rs9605903 | 100.00000 | 0.1166667 | 0 | | |
| 22_15447504_rs5747968 | 100.00000 | 0.1583333 | 1 | | |
| 22_15452483_rs2236639 | 100.00000 | 0.0625000 | 0 | | |

Check Mendelian errors:

```
> cMend <- checkMendelian(hm.a2, quiet = FALSE)
```

```
converting geneSet object to numerical matrix...
checking Mendelian errors ...
checking compatibility ...
```

Number of Mendelian errors per marker:

```
> head(cMend$nMerrMarker)
```

| | | | |
|-----------------------|-----------------------|-----------------------|-----------------------|
| 22_14884399_rs4911642 | 22_15298335_rs2027653 | 22_15412698_rs5747620 | 22_15434720_rs9605903 |
| 8 | 1 | 0 | 0 |
| 22_15447504_rs5747968 | 22_15452483_rs2236639 | | |
| 0 | 0 | | |

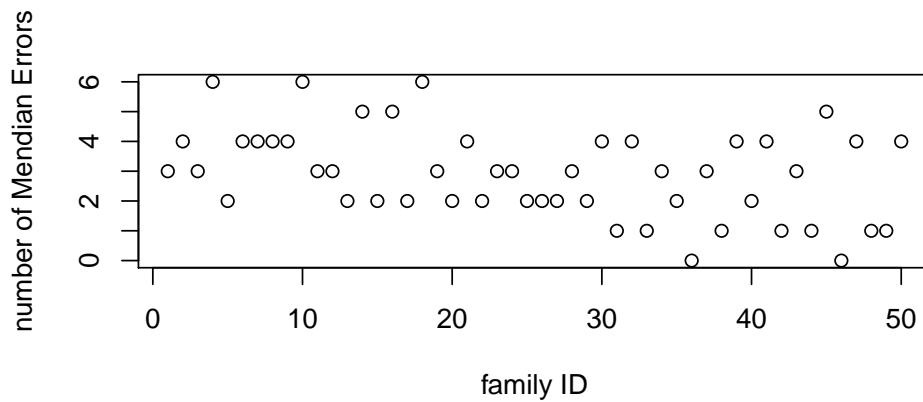
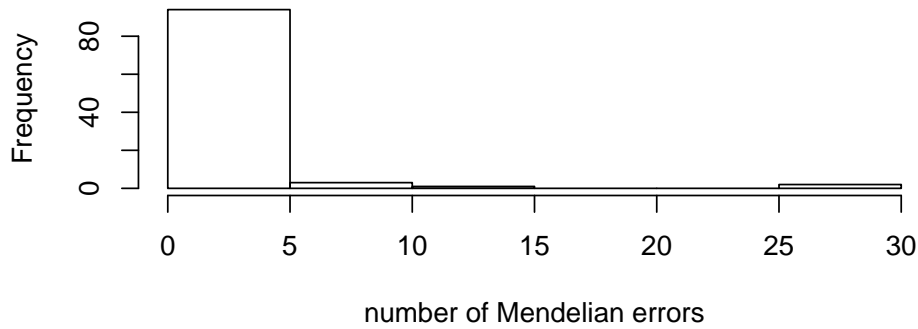
Number of Mendelian errors per family:

```
> head(cMend$nMerrFamily)
```

| family1 | family2 | family3 | family4 | family5 | family6 |
|---------|---------|---------|---------|---------|---------|
| 3 | 4 | 3 | 6 | 2 | 4 |

Plot counts of Mendelian errors

```
> par(mfrow = c(2, 1))
> hist(cMend$nMerrMarker, main = "", xlab = "number of Mendelian errors")
> plot(1:length(cMend$nMerrFamily), cMend$nMerrFamily, xlab = "family ID",
+      ylab = "number of Mendian Errors")
> par(mfrow = c(1, 1))
```



5.1 Summary of allele/genotype frequency based on GeneticsBase functions

Allele summary, including allele counts, allele frequencies, 95% CI of allele frequencies:

```
> t1 <- alleleSummary(hm.a2[1:10])
> t1
```

| Gene Marker | Position | Group | Allele | Count | Freq | CI-Lower | CI-Upper |
|---------------------------|----------|-------|--------|-------|-------|----------|----------|
| ALL 22_14884399_rs4911642 | ? | ALL | 2 | 48 | 0.151 | 0.113 | 0.192 |
| | | ALL | 4 | 270 | 0.849 | 0.808 | 0.887 |
| 22_15298335_rs2027653 | ? | ALL | 2 | 94 | 0.264 | 0.219 | 0.312 |
| | | ALL | 4 | 262 | 0.736 | 0.688 | 0.781 |
| 22_15412698_rs5747620 | ? | ALL | 2 | 169 | 0.469 | 0.417 | 0.522 |
| | | ALL | 4 | 191 | 0.531 | 0.478 | 0.583 |
| 22_15434720_rs9605903 | ? | ALL | 2 | 43 | 0.119 | 0.086 | 0.153 |
| | | ALL | 4 | 317 | 0.881 | 0.847 | 0.914 |

| | | | | | | | |
|------------------------|---|-----|---|-----|-------|-------|-------|
| 22_15447504_rs5747968 | ? | ALL | 3 | 59 | 0.164 | 0.128 | 0.203 |
| | | ALL | 4 | 301 | 0.836 | 0.797 | 0.872 |
| 22_15452483_rs2236639 | ? | ALL | 1 | 26 | 0.072 | 0.047 | 0.100 |
| | | ALL | 3 | 334 | 0.928 | 0.900 | 0.953 |
| 22_15455353_rs5747999 | ? | ALL | 1 | 225 | 0.625 | 0.575 | 0.675 |
| | | ALL | 2 | 135 | 0.375 | 0.325 | 0.425 |
| 22_15467656_rs11089263 | ? | ALL | 1 | 219 | 0.608 | 0.558 | 0.658 |
| | | ALL | 2 | 141 | 0.392 | 0.342 | 0.442 |
| 22_15474749_rs2096537 | ? | ALL | 1 | 80 | 0.231 | 0.188 | 0.277 |
| | | ALL | 2 | 266 | 0.769 | 0.723 | 0.812 |
| 22_15479107_rs9604959 | ? | ALL | 2 | 249 | 0.808 | 0.763 | 0.851 |
| | | ALL | 4 | 59 | 0.192 | 0.149 | 0.237 |

Footer:

Confidence intervals width is 95%, computed using the exact quantiles for the binomial distribution.

Same for genotypes:

```
> t2 <- genotypeSummary(hm.a2[1:10], founderOnly = TRUE)
> t2
```

| Gene Marker | Position | Group | Genotype | Count | Freq | CI-Lower | CI-Upper |
|-------------------------|----------|-------|----------|-------|-------|----------|----------|
| ? 22_14884399_rs4911642 | ? | ALL | 2/2 | 0 | 0.000 | | |
| | | | 2/4 | 33 | 0.308 | 0.224 | 0.402 |
| | | | 4/4 | 74 | 0.692 | 0.598 | 0.776 |
| | | | NA | 13 | | | |
| ? 22_15298335_rs2027653 | ? | ALL | 2/2 | 11 | 0.092 | 0.042 | 0.151 |
| | | | 2/4 | 41 | 0.345 | 0.261 | 0.429 |
| | | | 4/4 | 67 | 0.563 | 0.471 | 0.655 |
| | | | NA | 1 | | | |
| ? 22_15412698_rs5747620 | ? | ALL | 2/2 | 28 | 0.233 | 0.158 | 0.308 |
| | | | 2/4 | 60 | 0.500 | 0.408 | 0.592 |
| | | | 4/4 | 32 | 0.267 | 0.192 | 0.350 |
| | | | NA | 0 | | | |
| ? 22_15434720_rs9605903 | ? | ALL | 2/2 | 4 | 0.033 | 0.008 | 0.067 |
| | | | 2/4 | 20 | 0.167 | 0.100 | 0.233 |
| | | | 4/4 | 96 | 0.800 | 0.725 | 0.867 |
| | | | NA | 0 | | | |
| ? 22_15447504_rs5747968 | ? | ALL | 3/3 | 5 | 0.042 | 0.008 | 0.083 |
| | | | 3/4 | 28 | 0.233 | 0.158 | 0.308 |
| | | | 4/4 | 87 | 0.725 | 0.642 | 0.800 |
| | | | NA | 0 | | | |

| | | | | | | | | |
|---|------------------------|---|-----|-----|-----|-------|-------|-------|
| ? | 22_15452483_rs2236639 | ? | ALL | 1/1 | 2 | 0.017 | 0.000 | 0.042 |
| | | | | 1/3 | 11 | 0.092 | 0.042 | 0.150 |
| | | | | 3/3 | 107 | 0.892 | 0.833 | 0.942 |
| | | | | NA | 0 | | | |
| ? | 22_15455353_rs5747999 | ? | ALL | 1/1 | 49 | 0.408 | 0.325 | 0.500 |
| | | | | 1/2 | 54 | 0.450 | 0.358 | 0.542 |
| | | | | 2/2 | 17 | 0.142 | 0.083 | 0.208 |
| | | | | NA | 0 | | | |
| ? | 22_15467656_rs11089263 | ? | ALL | 1/1 | 46 | 0.383 | 0.300 | 0.475 |
| | | | | 1/2 | 52 | 0.433 | 0.342 | 0.525 |
| | | | | 2/2 | 22 | 0.183 | 0.117 | 0.258 |
| | | | | NA | 0 | | | |
| ? | 22_15474749_rs2096537 | ? | ALL | 1/1 | 0 | 0.000 | | |
| | | | | 1/2 | 51 | 0.440 | 0.353 | 0.534 |
| | | | | 2/2 | 65 | 0.560 | 0.466 | 0.647 |
| | | | | NA | 4 | | | |
| ? | 22_15479107_rs9604959 | ? | ALL | 2/2 | 68 | 0.642 | 0.547 | 0.736 |
| | | | | 2/4 | 36 | 0.340 | 0.255 | 0.434 |
| | | | | 4/4 | 2 | 0.019 | 0.000 | 0.047 |
| | | | | NA | 14 | | | |

| Expected | Obs-Exp | HWE | X ² | P-value |
|----------|---------|-------|----------------|---------|
| 2.544 | -2.544 | 3.557 | 0.0651 | |
| 27.911 | 5.089 | | | |
| 76.544 | -2.544 | | | |

| | | | |
|--------|--------|-------|-------|
| 8.338 | 2.662 | 1.572 | 0.241 |
| 46.324 | -5.324 | | |
| 64.338 | 2.662 | | |

| | | | |
|--------|--------|-------|---|
| 28.033 | -0.033 | 0.000 | 1 |
| 59.933 | 0.067 | | |
| 32.033 | -0.033 | | |

| | | | |
|--------|--------|-------|--------|
| 1.633 | 2.367 | 4.395 | 0.0633 |
| 24.733 | -4.733 | | |
| 93.633 | 2.367 | | |

| | | | |
|--------|--------|-------|-------|
| 3.008 | 1.992 | 1.861 | 0.305 |
| 31.983 | -3.983 | | |
| 85.008 | 1.992 | | |

```
0.469    1.531    5.691    0.0501
14.062   -3.062
105.469  1.531
```

```
48.133   0.867   0.116   0.84
55.733   -1.733
16.133   0.867
```

```
43.200   2.800   1.134   0.344
57.600   -5.600
19.200   2.800
```

```
5.606    -5.606   9.210   0.003
39.789   11.211
70.606   -5.606
```

```
69.774   -1.774   1.266   0.343
32.453   3.547
3.774    -1.774
```

Footer:

Confidence intervals width is 95%, computed using the exact quantiles for the binomial distribution. As the true distribution is multinomial this is only approximately correct.

HWE test:

```
> hwe <- HWE(hm.a2[1:10])
> hwe
```

```
[[1]]
[1] "diseq"
```

```
$call
HWE(object = hm.a2[1:10])
```

```
$D
              Est      2.5%      97.5%    n      P-value
22_14884399_rs4911642  0.022783909  0.01353783  0.035599858 159 2.602556e-02
22_15298335_rs2027653 -0.025785886 -0.05535128  0.005057758 178 8.267860e-02
22_15412698_rs5747620 -0.001844136 -0.03876543  0.033456790 180 1.000000e+00
22_15434720_rs9605903 -0.019066358 -0.04258488  0.001859568 180 2.387117e-02
22_15447504_rs5747968 -0.023140432 -0.04684414  0.001111111 180 2.881018e-02
22_15452483_rs2236639 -0.005895062 -0.02128858  0.006049383 180 2.254611e-01
22_15455353_rs5747999  0.012847222 -0.01987654  0.045679012 180 5.265022e-01
22_15467656_rs11089263 -0.018819444 -0.05332562  0.017777778 180 2.780714e-01
```

```

22_15474749_rs2096537 0.053459855 0.03749708 0.072245982 173 7.796707e-06
22_15479107_rs9604959 0.010720611 -0.01239669 0.034333361 154 6.010626e-01

```

\$`D`

| | Est | 2.5% | 97.5% | n | P-value |
|------------------------|--------------|------------|-------------|-----|--------------|
| 22_14884399_rs4911642 | 0.177777778 | 0.1316726 | 0.232558140 | 159 | 2.602556e-02 |
| 22_15298335_rs2027653 | -0.369850611 | -0.7800000 | 0.024598028 | 178 | 8.267860e-02 |
| 22_15412698_rs5747620 | -0.008368054 | -0.1812500 | 0.133893281 | 180 | 1.000000e+00 |
| 22_15434720_rs9605903 | -1.336398053 | -2.7869822 | 0.018426485 | 180 | 2.387117e-02 |
| 22_15447504_rs5747968 | -0.861534042 | -1.7551020 | 0.006944444 | 180 | 2.881018e-02 |
| 22_15452483_rs2236639 | -1.130177515 | -4.4442344 | 0.084337349 | 180 | 2.254611e-01 |
| 22_15455353_rs5747999 | 0.054814815 | -0.1337868 | 0.197900587 | 180 | 5.265022e-01 |
| 22_15467656_rs11089263 | -0.122679946 | -0.3591837 | 0.073075837 | 180 | 2.780714e-01 |
| 22_15474749_rs2096537 | 0.300751880 | 0.2401434 | 0.367588933 | 173 | 7.796707e-06 |
| 22_15479107_rs9604959 | 0.069226057 | -0.3750000 | 0.201709296 | 154 | 6.010626e-01 |

\$r

| | Est | 2.5% | 97.5% | n | P-value |
|------------------------|--------------|--------------|-------------|-----|--------------|
| 22_14884399_rs4911642 | -0.177777778 | -0.232558140 | -0.13167260 | 159 | 2.602556e-02 |
| 22_15298335_rs2027653 | 0.132694494 | -0.024598028 | 0.27724541 | 178 | 8.267860e-02 |
| 22_15412698_rs5747620 | 0.007404195 | -0.133893281 | 0.15513834 | 180 | 1.000000e+00 |
| 22_15434720_rs9605903 | 0.181277969 | -0.018426485 | 0.36507937 | 180 | 2.387117e-02 |
| 22_15447504_rs5747968 | 0.168872121 | -0.006944444 | 0.33751894 | 180 | 2.881018e-02 |
| 22_15452483_rs2236639 | 0.087977890 | -0.084337349 | 0.29941618 | 180 | 2.254611e-01 |
| 22_15455353_rs5747999 | -0.054814815 | -0.197900587 | 0.08210097 | 180 | 5.265022e-01 |
| 22_15467656_rs11089263 | 0.078985718 | -0.073075837 | 0.22285714 | 180 | 2.780714e-01 |
| 22_15474749_rs2096537 | -0.300751880 | -0.367588933 | -0.24014337 | 173 | 7.796707e-06 |
| 22_15479107_rs9604959 | -0.069226057 | -0.201709296 | 0.08333333 | 154 | 6.010626e-01 |

\$`X^2`

| | Est | 2.5% | 97.5% | n | P-value |
|------------------------|--------------|------|-------|-----|--------------|
| 22_14884399_rs4911642 | 0.0316049383 | NA | NA | 159 | 2.602556e-02 |
| 22_15298335_rs2027653 | 0.0176078288 | NA | NA | 178 | 8.267860e-02 |
| 22_15412698_rs5747620 | 0.0000548221 | NA | NA | 180 | 1.000000e+00 |
| 22_15434720_rs9605903 | 0.0328617022 | NA | NA | 180 | 2.387117e-02 |
| 22_15447504_rs5747968 | 0.0285177933 | NA | NA | 180 | 2.881018e-02 |
| 22_15452483_rs2236639 | 0.0077401092 | NA | NA | 180 | 2.254611e-01 |
| 22_15455353_rs5747999 | 0.0030046639 | NA | NA | 180 | 5.265022e-01 |
| 22_15467656_rs11089263 | 0.0062387437 | NA | NA | 180 | 2.780714e-01 |
| 22_15474749_rs2096537 | 0.0904516931 | NA | NA | 173 | 7.796707e-06 |
| 22_15479107_rs9604959 | 0.0047922469 | NA | NA | 154 | 6.010626e-01 |

Defatult graphical display of LD:

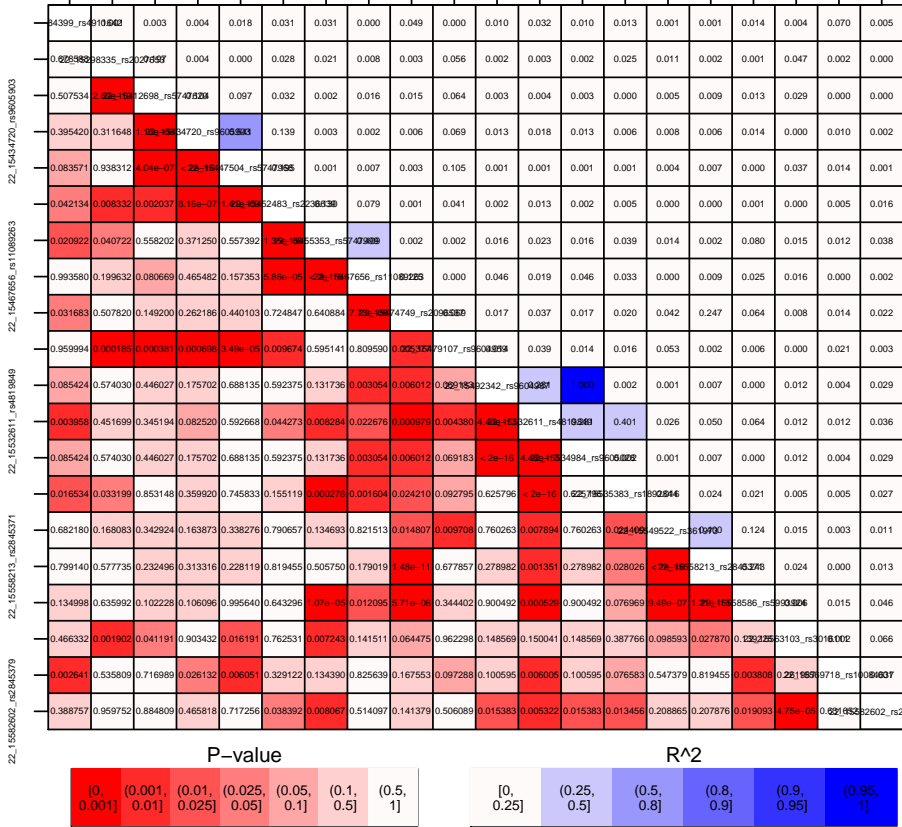
```

> ld.small <- LD(hm.a2[1:20])
> plot(ld.small)

```

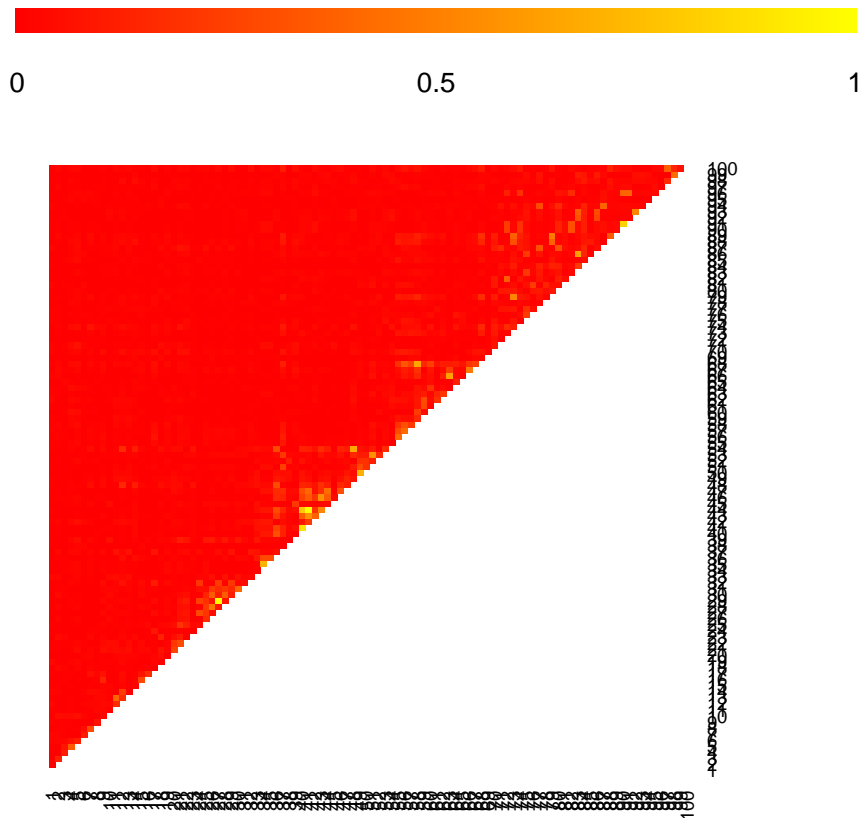
Linkage Disequilibrium

22_14884399_rs4911642 22_15434720_rs9605903 22_15455353_rs5747999 22_15479107_rs9604959 22_15534984_rs9605028 22_15558213_rs2845371 22_15569718_rs10084637



Alternative graphical display of LD:

```
> ld <- LD(hm.a2)
> LDView(ld@"X^2")
```



6 Hypothesis Testing

6.1 Armitage test

For the following examples, suppose 'A' is the minor allele, and 'a' is the major allele.

Armitage test using additive model to code genotype:

| genotype | coding |
|----------|--------|
| AA | 2 |
| Aa | 1 |
| aa | 0 |

```
> res.A <- Armitage(hm.a2, method = "A")
> head(res.A)
```

| | stat | pvalue |
|-----------------------|-------------|--------------|
| 22_14884399_rs4911642 | 4.38726430 | 3.620837e-02 |
| 22_15298335_rs2027653 | 8.62689991 | 3.312348e-03 |
| 22_15412698_rs5747620 | 0.01107079 | 9.162030e-01 |
| 22_15434720_rs9605903 | 34.00571358 | 5.495048e-09 |
| 22_15447504_rs5747968 | 35.11899027 | 3.101613e-09 |
| 22_15452483_rs2236639 | 2.43861135 | 1.183810e-01 |

| | genotype | coding |
|---|----------|--------|
| Armitage test using recessive model to code genotype: | AA | 1 |
| | Aa | 0 |
| | aa | 0 |

```
> res.R <- Armitage(hm.a2, method = "R")
> head(res.R)
```

| | stat | pvalue |
|-----------------------|-----------|-------------|
| 22_14884399_rs4911642 | NA | NA |
| 22_15298335_rs2027653 | 1.6258677 | 0.202275552 |
| 22_15412698_rs5747620 | 0.1285714 | 0.719917853 |
| 22_15434720_rs9605903 | 6.2068966 | 0.012725353 |
| 22_15447504_rs5747968 | 9.4736842 | 0.002084403 |
| 22_15452483_rs2236639 | 0.0000000 | 1.000000000 |

| | genotype | coding |
|--|----------|--------|
| Armitage test using dominant model to code genotype: | AA | 1 |
| | Aa | 1 |
| | aa | 0 |

```
> res.D <- Armitage(hm.a2, method = "D")
> head(res.D)
```

| | stat | pvalue |
|-----------------------|------------|--------------|
| 22_14884399_rs4911642 | 4.3872643 | 3.620837e-02 |
| 22_15298335_rs2027653 | 10.0936094 | 1.487844e-03 |
| 22_15412698_rs5747620 | 0.2462380 | 6.197365e-01 |
| 22_15434720_rs9605903 | 37.0478170 | 1.152676e-09 |
| 22_15447504_rs5747968 | 35.8892308 | 2.088600e-09 |
| 22_15452483_rs2236639 | 3.0769231 | 7.941063e-02 |

6.2 Logistic regression

First, we need to construct some synthetic covariates on the founders

```
> sampleInfo(hm.f)$race <- sampleInfo(hm.f)$affected
> raceval <- sampleInfo(hm.f)$race - 1
> sampleInfo(hm.f)$Norm0.0 <- rnorm(nobs(hm.f), mean = 0 * raceval)
> sampleInfo(hm.f)$Norm0.5 <- rnorm(nobs(hm.f), mean = 0.5 * raceval)
> sampleInfo(hm.f)$Norm1.0 <- rnorm(nobs(hm.f), mean = 1 * raceval)
> sampleInfo(hm.f)$Norm1.5 <- rnorm(nobs(hm.f), mean = 1.5 * sampleInfo(hm.f)$race)
> doSample <- function(raceval, mult) {
+   prob <- c(0.33 - raceval * mult, 0.33 + (raceval * mult)/2, 0.33 +
+     (raceval * mult)/2)
+   factor(sample(x = c("Red", "Green", "Blue"), size = 1, p = prob,
+     rep = T))
+ }
> sampleInfo(hm.f)$Cat0.0 <- sapply(raceval, doSample, mult = 0)
> sampleInfo(hm.f)$Cat0.1 <- sapply(raceval, doSample, mult = 0.1)
> sampleInfo(hm.f)$Cat0.2 <- sapply(raceval, doSample, mult = 0.2)
> sampleInfo(hm.f)$Cat0.3 <- sapply(raceval, doSample, mult = 0.3)
```

Now, construct a function to fit the regression model and return the parameters and statistics that are of interest.

```

model <- function( markerName )
{
  # extract requested genetic marker
  genotype <- genotypes(hm.f,marker=markerName)

  # get data frame to use for fitting the model
  mframe <- model.frame(race ~ sex + Norm0.0 + Norm0.5 + Norm1.0 + Norm1.5 + genotype,
                        data=sampleInfo(hm.f) )

  # To test significance of a term, best method is to do anova of
  # the full model against a submodel omitting the particular term.
  # This avoids issues with changes in names of factor levels,
  # presence or absence of covariates, etc.
  result <- try(
    {
fit.with    <- glm( race==1 ~ sex + Norm0.0 + Norm0.5 + Norm1.0 + Norm1.5 + as.factor(genotype),
                    data=mframe, family="binomial")
fit.without <- glm( race==1 ~ sex + Norm0.0 + Norm0.5 + Norm1.0 + Norm1.5,
                    data=mframe, family="binomial")
                    anova(fit.with, fit.without, test="Chisq")$"P(>|Chi|)"[2]
    }
  )

  if(class(result)=="try-error")
    return(NA) # or return(result) to see the error messages
  else

    result # full result. Usually we want to specify exactly which
           # parameters and stats get returned so the format is consistent
           # across all markers.
}

```

```

> model <- function(markerName) {
+   genotype <- genotypes(hm.f, marker = markerName)
+   mframe <- model.frame(race ~ sex + Norm0.0 + Norm0.5 + Norm1.0 +
+     Norm1.5 + genotype, data = sampleInfo(hm.f))
+   result <- try({
+     fit.with <- glm(race == 1 ~ sex + Norm0.0 + Norm0.5 + Norm1.0 +
+       Norm1.5 + as.factor(genotype), data = mframe, family = "binomial")
+     fit.without <- glm(race == 1 ~ sex + Norm0.0 + Norm0.5 + Norm1.0 +
+       Norm1.5, data = mframe, family = "binomial")
+     anova(fit.with, fit.without, test = "Chisq")$"P(>|Chi|)"[2]
+   })
+   if (class(result) == "try-error")
+     return(NA)
+   else result
+ }

```

Fit to a subset of 50, then 100 and use this information to compute the expected run time for all markers:

```

> t1 <- unix.time(fits <- sapply(markerNames(hm.f)[1:50], model))[3]
> t1

```

```
elapsed
3.554
```

(This takes 5.365 seconds on my MacBook Pro, R 2.4.1)

```
> t2 <- unix.time(fits <- sapply(markerNames(hm.f)[1:100], model))[3]
> t2
```

```
elapsed
6.767
```

(This takes 11.878 seconds on my MacBook Pro, R 2.4.1)

Estimate total time to complete, in minutes

```
> t1 + (t2 - t1)/50 * (nmarker(hm.f) - 50)/60
```

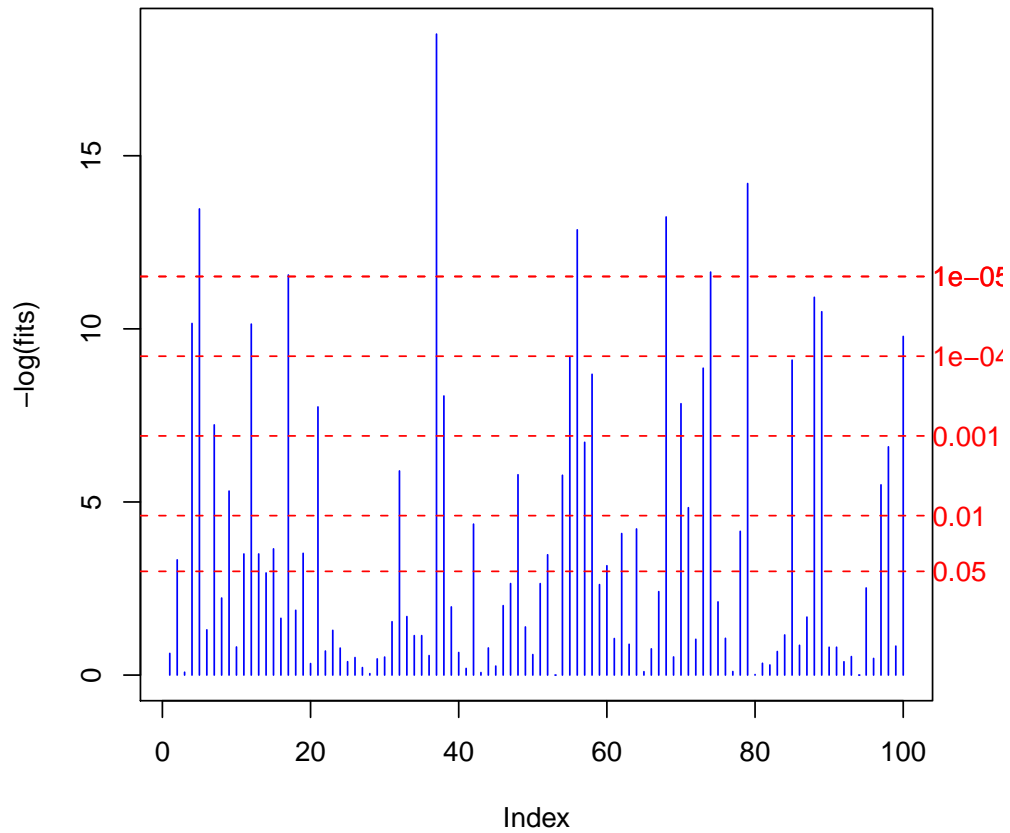
```
elapsed
12.26551
```

(This yields 23.02 minutes on my MacBook Pro, R 2.4.1)

Plot the p-values for the first 100 markers

```
> fits.sorted <- sort(fits)
> plot(-log(fits), type = "h", col = "blue")
> labels <- c(0.05, 0.01, 0.001, 1e-04, 1e-05, 1e-05)
> abline(h = -log(labels), lty = 2, col = "red")
> mtext(text = as.character(labels), side = 4, at = -log(labels), col = "red",
+       las = 1)
> title("Per-marker statistical significance")
```


Per-marker statistical significance



6.3 Family-Based Association Test ('fbat')

Do the fbat calculations:

```
> f <- fbat(hm.a2)
```

Show the p-values:

```
> summaryPvalue(f)
```

```
*****
              chisq rank    pvalue
22_14884399_rs4911642  1.6666667    1 0.19670560
22_15298335_rs2027653  0.0666667    1 0.79625341
22_15412698_rs5747620  0.57142857   1 0.44969180
22_15434720_rs9605903  1.0000000    1 0.31731051
22_15447504_rs5747968  1.0000000    1 0.31731051
22_15452483_rs2236639  0.0000000    1 1.00000000
22_15455353_rs5747999  1.05882353   1 0.30348366
22_15467656_rs11089263 4.5000000    1 0.03389485
22_15474749_rs2096537  0.75757576   1 0.38408825
```

| | | | |
|------------------------|------------|---|------------|
| 22_15479107_rs9604959 | 2.77777778 | 1 | 0.09558070 |
| 22_15492342_rs9604967 | 1.00000000 | 1 | 0.31731051 |
| 22_15532611_rs4819849 | 1.00000000 | 1 | 0.31731051 |
| 22_15534984_rs9605028 | 1.00000000 | 1 | 0.31731051 |
| 22_15535383_rs1892844 | 0.00000000 | 0 | 0.00000000 |
| 22_15549522_rs361973 | 0.80645161 | 1 | 0.36917142 |
| 22_15558213_rs2845371 | 0.04761905 | 1 | 0.82725935 |
| 22_15558586_rs5993924 | 0.20000000 | 1 | 0.65472085 |
| 22_15563103_rs3016111 | 0.00000000 | 0 | 0.00000000 |
| 22_15569718_rs10084637 | 0.69230769 | 1 | 0.40538056 |
| 22_15582602_rs2845379 | 4.16666667 | 1 | 0.04122683 |
| 22_15583103_rs2845380 | 0.00000000 | 0 | 0.00000000 |
| 22_15594252_rs2845346 | 0.00000000 | 1 | 1.00000000 |
| 22_15608796_rs17433377 | 0.00000000 | 0 | 0.00000000 |
| 22_15634399_rs2190742 | 1.08695652 | 1 | 0.29714653 |
| 22_15636231_rs5748614 | 3.76923077 | 1 | 0.05220364 |
| 22_15644565_rs5748622 | 3.57142857 | 1 | 0.05878172 |
| 22_15644904_rs9605145 | 3.57142857 | 1 | 0.05878172 |
| 22_15645124_rs5748623 | 3.60000000 | 1 | 0.05777957 |
| 22_15645194_rs9605146 | 5.14285714 | 1 | 0.02334220 |
| 22_15647006_rs759235 | 0.60000000 | 1 | 0.43857803 |
| 22_15649076_rs2108585 | 2.90909091 | 1 | 0.08808151 |
| 22_15653728_rs9606468 | 0.00000000 | 1 | 1.00000000 |
| 22_15655394_rs5748636 | 0.36000000 | 1 | 0.54850624 |
| 22_15658762_rs4819535 | 0.69230769 | 1 | 0.40538056 |
| 22_15660822_rs5748648 | 0.69230769 | 1 | 0.40538056 |
| 22_15661931_rs738045 | 0.60000000 | 1 | 0.43857803 |
| 22_15665949_rs2385714 | 0.00000000 | 0 | 0.00000000 |
| 22_15668988_rs2072467 | 0.00000000 | 0 | 0.00000000 |
| 22_15669118_rs2072466 | 1.00000000 | 1 | 0.31731051 |
| 22_15674251_rs7291429 | 0.00000000 | 1 | 1.00000000 |
| 22_15681217_rs874835 | 0.03703704 | 1 | 0.84738966 |
| 22_15681843_rs874836 | 2.46153846 | 1 | 0.11666446 |
| 22_15684246_rs175139 | 2.46153846 | 1 | 0.11666446 |
| 22_15684887_rs983305 | 0.03703704 | 1 | 0.84738966 |
| 22_15686104_rs175140 | 0.66666667 | 1 | 0.41421618 |
| 22_15690741_rs175149 | 1.28571429 | 1 | 0.25683926 |
| 22_15692596_rs9606481 | 0.61538462 | 1 | 0.43276758 |
| 22_15695102_rs17363716 | 0.00000000 | 0 | 0.00000000 |
| 22_15695503_rs165757 | 0.11764706 | 1 | 0.73160059 |
| 22_15697233_rs175152 | 0.80645161 | 1 | 0.36917142 |
| 22_15698150_rs165611 | 0.92592593 | 1 | 0.33592381 |
| 22_15699156_rs165778 | 1.00000000 | 1 | 0.31731051 |
| 22_15706181_rs165810 | 0.00000000 | 0 | 0.00000000 |
| 22_15706432_rs2075120 | 0.00000000 | 0 | 0.00000000 |
| 22_15775610_rs737936 | 1.00000000 | 1 | 0.31731051 |
| 22_15776612_rs7288841 | 0.00000000 | 0 | 0.00000000 |
| 22_15777875_rs9606534 | 2.00000000 | 1 | 0.15729921 |
| 22_15778508_rs7292561 | 3.00000000 | 1 | 0.08326452 |
| 22_15778800_rs7293026 | 0.80645161 | 1 | 0.36917142 |
| 22_15778812_rs13058496 | 3.00000000 | 1 | 0.08326452 |

| | | | |
|------------------------|------------|---|------------|
| 22_15779211_rs8136206 | 0.80000000 | 1 | 0.37109337 |
| 22_15785173_rs759081 | 1.00000000 | 1 | 0.31731051 |
| 22_15787349_rs5992587 | 1.80000000 | 1 | 0.17971249 |
| 22_15787566_rs11703901 | 2.66666667 | 1 | 0.10247043 |
| 22_15789897_rs12485066 | 3.00000000 | 1 | 0.08326452 |
| 22_15790373_rs5748744 | 3.85714286 | 1 | 0.04953461 |
| 22_15791899_rs9306242 | 3.24000000 | 1 | 0.07186064 |
| 22_15792216_rs9605179 | 4.00000000 | 1 | 0.04550026 |
| 22_15792806_rs5992590 | 0.61538462 | 1 | 0.43276758 |
| 22_15794103_rs5994097 | 2.13043478 | 1 | 0.14439979 |
| 22_15794640_rs9618937 | 0.50000000 | 1 | 0.47950012 |
| 22_15795572_rs5748748 | 0.69230769 | 1 | 0.40538056 |
| 22_15806401_rs5748755 | 2.66666667 | 1 | 0.10247043 |
| 22_15807037_rs2385785 | 2.00000000 | 1 | 0.15729921 |
| 22_15809384_rs1981707 | 5.53846154 | 1 | 0.01860293 |
| 22_15809434_rs1981708 | 1.60000000 | 1 | 0.20590321 |
| 22_15813210_rs4819923 | 2.46153846 | 1 | 0.11666446 |
| 22_15813888_rs5994105 | 0.40000000 | 1 | 0.52708926 |
| 22_15814084_rs5748760 | 1.00000000 | 1 | 0.31731051 |
| 22_15816846_rs2385786 | 0.12500000 | 1 | 0.72367361 |
| 22_15821524_rs5994110 | 1.00000000 | 1 | 0.31731051 |
| 22_15822154_rs17733785 | 0.33333333 | 1 | 0.56370286 |
| 22_15823131_rs7287116 | 0.20000000 | 1 | 0.65472085 |
| 22_15825502_rs5748765 | 0.03030303 | 1 | 0.86180443 |
| 22_15826157_rs1541529 | 0.00000000 | 0 | 0.00000000 |
| 22_15826914_rs5748766 | 0.81818182 | 1 | 0.36571230 |
| 22_15830515_rs2041607 | 0.69230769 | 1 | 0.40538056 |
| 22_15832966_rs757630 | 0.00000000 | 0 | 0.00000000 |
| 22_15842185_rs4819932 | 1.00000000 | 1 | 0.31731051 |
| 22_15847411_rs4819934 | 0.22222222 | 1 | 0.63735189 |
| 22_15847684_rs4819936 | 0.22222222 | 1 | 0.63735189 |
| 22_15850779_rs9618954 | 1.60000000 | 1 | 0.20590321 |
| 22_15855921_rs2399152 | 2.27272727 | 1 | 0.13166802 |
| 22_15868195_rs5748798 | 0.00000000 | 0 | 0.00000000 |
| 22_15869577_rs7291404 | 0.06666667 | 1 | 0.79625341 |
| 22_15869890_rs11913227 | 0.18181818 | 1 | 0.66981536 |
| 22_15870932_rs5994128 | 0.14285714 | 1 | 0.70545699 |
| 22_15872203_rs5994129 | 1.50000000 | 1 | 0.22067136 |
| 22_15872452_rs917838 | 0.18181818 | 1 | 0.66981536 |
| 22_15872533_rs2192155 | 0.00000000 | 0 | 0.00000000 |

Look at the fit details for a specific marker:

```
> viewstat(f, "22_14884399_rs4911642")
```

```
50 pedigree 180 persons
13 informative families at marker 22_14884399_rs4911642
The alleles of marker 22_14884399_rs4911642 >>
[1] 1 2
Score for marker 22_14884399_rs4911642 >>
[1] 5 21
```

```

Expected score for marker 22_14884399_rs4911642 >>
[1] 7.5 18.5
Covariance matrix of the score for marker 22_14884399_rs4911642 >>
      [,1] [,2]
[1,] 3.75 -3.75
[2,] -3.75 3.75
Moore-Penrose generalized inverse of covariance matrix
      [,1] [,2]
[1,] 0.06666667 -0.06666667
[2,] -0.06666667 0.06666667
test statistics for marker 22_14884399_rs4911642 >>
      chisq      rank      pvalue
1.6666667 1.0000000 0.1967056
*****

```

7 Study planning tools (GeneticsDesign package)

7.1 Power to detect a low-frequency allele

Compute the probability of missing an allele with frequency 0.15 when 20 genotypes are sampled:

```
> gregorius(freq = 0.15, N = 20)
```

```
$call
```

```
gregorius(freq = 0.15, N = 20)
```

```
$method
```

```
[1] "Compute missprob given N and freq"
```

```
$freq
```

```
[1] 0.15
```

```
$N
```

```
[1] 20
```

```
$missprob
```

```
[1] 0.1938351
```

Determine what sample size is required to observe all alleles with true frequency 0.15 with probability 0.95

```
> gregorius(freq = 0.15, missprob = 1 - 0.95)
```

```
$call
```

```
gregorius(freq = 0.15, missprob = 1 - 0.95)
```

```
$method
```

```
[1] "Determine minimal N given missprob and freq"
```

```
$freq
```

```
[1] 0.15
```

```
$N
```

```
[1] 29
```

```
$missprob
```

```
[1] 0.04520557
```

7.2 Power for a genetics study using a quantitative outcome

Calculate power for genetics study using a quantitative outcome:

```
> GeneticPower.Quantitative.Numeric(N = 50, freq = 0.1, minh = "recessive",  
+   alpha = 0.05)
```

```
[1] 0.0600883
```

```
> GeneticPower.Quantitative.Factor(N = 50, freq = 0.1, minh = "recessive",  
+   alpha = 0.05)
```

```
[1] 0.08666032
```

For a range of sample sizes:

```
> power.range <- function(N, ...) {  
+   sapply(N, function(n) GeneticPower.Quantitative.Numeric(N = n, ...))  
+ }  
> power.range(N = c(25, 50, 100, 200, 500), freq = 0.1, minh = "recessive",  
+   alpha = 0.05)
```

```
[1] 0.05492509 0.06008830 0.07049383 0.09160716 0.15665828
```

Create a power table:

```
> fun <- function(p) power.range(freq = p, N = seq(100, 1000, by = 100),  
+   alpha = 0.05, minh = "recessive")  
> m <- sapply(X = seq(0.1, 0.9, by = 0.1), fun)  
> colnames(m) <- seq(0.1, 0.9, by = 0.1)  
> rownames(m) <- seq(100, 1000, by = 100)  
> print(round(m, 2))
```

| | 0.1 | 0.2 | 0.3 | 0.4 | 0.5 | 0.6 | 0.7 | 0.8 | 0.9 |
|------|------|------|------|------|------|------|------|------|------|
| 100 | 0.07 | 0.20 | 0.47 | 0.76 | 0.93 | 0.98 | 0.99 | 0.99 | 0.97 |
| 200 | 0.09 | 0.35 | 0.77 | 0.97 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 |
| 300 | 0.11 | 0.49 | 0.91 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 |
| 400 | 0.13 | 0.61 | 0.97 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 |
| 500 | 0.16 | 0.70 | 0.99 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 |
| 600 | 0.18 | 0.78 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 |
| 700 | 0.20 | 0.84 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 |
| 800 | 0.22 | 0.88 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 |
| 900 | 0.24 | 0.92 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 |
| 1000 | 0.27 | 0.94 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 | 1.00 |

7.3 Power calculator for genetic linear trend association studies.

The power is for the test that disease is associated with a marker, given high risk allele frequency ('A'), disease prevalence, genotype relative risk ('Aa'), genotype relative risk ('AA'), LD measure (D' or R^2), marker allele frequency ('B'), number of cases, control:case ratio, and probability of the Type I error. The linear trend test (Cochran 1954; Armitage 1955) is used.

Using R^2 as the measuer of LD:

```
> res1 <- GPC(pA = 0.05, pD = 0.1, RRAa = 1.414, RRAA = 2, r2 = 0.9, pB = 0.06,
+ nCase = 500, ratio = 1, alpha = 0.05, quiet = FALSE)
```

```
Case-control parameters>>
```

```

                                     [,1]
Number of cases                      500.00000000
Number of controls                   500.00000000
High risk allele frequency (A)       0.05000000
Prevalence                           0.10000000
Genotypic relative risk Aa           1.41400000
Genotypic relative risk AA           2.00000000
Genotypic risk for aa (baseline)     0.09598495
```

```
Marker locus B>>
```

```

                                     [,1]
High risk allele frequency (B)       0.06000000
Linkage disequilibrium (D')          0.99723021
Penetrance at marker genotype bb     0.09599596
Penetrance at marker genotype Bb     0.12902094
Penetrance at marker genotype BB     0.17344738
Genotypic odds ratio Bb              1.39498627
Genotypic odds ratio BB              1.97612611
```

```
Expected allele frequencies>>
```

```

      Case   Control
B 0.07901192 0.05788756
b 0.92098808 0.94211244
```

```
Expected genotype frequencies>>
```

```

      Case   Control
BB 0.006244106 0.003306210
Bb 0.145535624 0.109162708
bb 0.848220271 0.887531081
```

```
Case-control statistics>>
```

```

Alpha   Power
0.100 0.58900688
0.050 0.46393515
0.010 0.23992694
0.001 0.07762229
0.050 0.46393515
```

```
power (alpha= 0.05 )= 0.4639352 ncp= 3.494199
```

```
Using  $D'$  as the measure of LD:
```

```
> res2 <- GPC.default(pA = 0.05, pD = 0.1, RRAa = 1.414, RRAA = 2, Dprime = 0.9,
+ pB = 0.06, nCase = 500, ratio = 1, alpha = 0.05, quiet = FALSE)
```

```
Case-control parameters>>
```

```

                                     [,1]
Number of cases                      500.00000000
Number of controls                   500.00000000
```

High risk allele frequency (A) 0.05000000
Prevalence 0.10000000
Genotypic relative risk Aa 1.41400000
Genotypic relative risk AA 2.00000000
Genotypic risk for aa (baseline) 0.09598495

Marker locus B>>

[,1]
High risk allele frequency (B) 0.06000000
Linkage disequilibrium (D') 0.90000000
Penetrance at marker genotype bb 0.09638274
Penetrance at marker genotype Bb 0.12624798
Penetrance at marker genotype BB 0.16539976
Genotypic odds ratio Bb 1.35463248
Genotypic odds ratio BB 1.85798248

Expected allele frequencies>>

| | Case | Control |
|---|------------|------------|
| B | 0.07715825 | 0.05809353 |
| b | 0.92284175 | 0.94190647 |

Expected genotype frequencies>>

| | Case | Control |
|----|-------------|-------------|
| BB | 0.005954391 | 0.003338401 |
| Bb | 0.142407718 | 0.109510254 |
| bb | 0.851637891 | 0.887151345 |

Case-control statistics>>

| Alpha | Power |
|-------|------------|
| 0.100 | 0.52110203 |
| 0.050 | 0.39631201 |
| 0.010 | 0.18968278 |
| 0.001 | 0.05549084 |
| 0.050 | 0.39631201 |

power (alpha= 0.05)= 0.396312 ncp= 2.878885

The End.