

An Integrated Genetic Analysis Package Using R

Jing Hua Zhao

MRC Epidemiology Unit, Cambridge, UK
<http://www.mrc-epid.cam.ac.uk>

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1 Introduction

This package was initiated to integrate some C/Fortran/SAS programs I have written or used over the years. As such, it would rather be a long-term project, but an immediate benefit would be something complementary to other packages currently available from CRAN, e.g. **genetics**, **hwde**, etc. I hope eventually this will be part of a bigger effort to fulfill most of the requirements foreseen by many, e.g. Guo and Lange (2000), within the portable environment of R for data management, analysis, graphics and object-oriented programming. My view has been outlined more formally in Zhao and Tan (2006b) and Zhao and Tan (2006a) in relation to other package systems. Also reported are Zhao (2005) and Zhao (2006) on package **kinship**.

The number of functions are quite limited and experimental, but I already feel the enormous advantage by shifting to R and would like sooner rather than later to share my work with others. I will not claim this work as exclusively done by me, but would like to invite others to join me and enlarge the collections and improve them.

2 Implementation

The following list shows the data and functions currently available.

| | |
|------|--------------------------------------|
| BFDP | Bayesian false-discovery probability |
| FPRP | False-positive report probability |

| | |
|----------------|--|
| aldh2 | ALDH2 markers and Alcoholism |
| apoeapoc | APOE/APOC1 markers and Schizophrenia |
| bt | Bradley-Terry model for contingency table |
| ccsize | Power and sample size for case-cohort design |
| chow.test | Chow's test for heterogeneity in two regressions |
| cf | Cystic Fibrosis data |
| comp.score | score statistics for testing genetic linkage of quantitative trait |
| crohn | Crohn disease data |
| fa | Friedreich Ataxia data |
| fbsize | Sample size for family-based linkage and association design |
| fsnps | A case-control data involving four SNPs with missing genotype |
| gc.em | Gene counting for haplotype analysis |
| gcontrol | genomic control |
| gcp | Permutation tests using GENECOUNTING |
| genecounting | Gene counting for haplotype analysis |
| gif | Kinship coefficient and genetic index of familiarity |
| hap | Haplotype reconstruction |
| hap.em | Gene counting for haplotype analysis |
| hap.score | Score Statistics for Association of Traits with Haplotypes |
| hla | HLA markers and Schizophrenia |
| htr | Haplotype trend regression |
| hwe | Hardy-Weinberg equilibrium test for multiallelic marker |
| hwe.hardy | Hardy-Weinberg equilibrium test using MCMC |
| kbyl | LD statistics for two multiallelic loci |
| kin.morgan | kinship matrix for simple pedigree |
| makeped | A function to prepare pedigrees in post-MAKEPED format |
| mao | A study of Parkinson's disease and MAO gene |
| mia | multiple imputation analysis for hap |
| mtdt | Transmission/disequilibrium test of a multiallelic marker |
| muvar | Means and variances under 1- and 2- locus (biallelic) QTL model |
| nep499 | A study of Alzheimer's disease with eight SNPs and APOE |
| pbsize | Power for population-based association design |
| pedtodot | Converting pedigree(s) to dot file(s) |
| pfc | Probability of familial clustering of disease |
| pfc.sim | Probability of familial clustering of disease |
| pgc | Preparing weight for GENECOUNTING |
| plot.hap.score | Plot Haplotype Frequencies versus Haplotype |

| | |
|------------------------------|---|
| | Score Statistics |
| <code>print.hap.score</code> | Print a <code>hap.score</code> object |
| <code>qqunif</code> | Q-Q plot for uniformly distributed random variable |
| <code>s2k</code> | Statistics for 2 by K table |
| <code>snca</code> | A study of Parkinson's disease and SNCA makers |
| <code>tbyt</code> | LD statistics for two SNPs |
| <code>tsc</code> | Power calculation for two-stage case-control design |
| <code>twinan90</code> | Classic twin models |
| <code>whscore</code> | Whittemore-Halpern scores for allele-sharing |

Assuming proper installation, you will be able to obtain the list by typing `library(help=gap)` or view the list within a web browser via `help.start()`. A PDF version of this file can be viewed with command `vignette("gap",package="gap")`.

You can cut and paste examples at end of each function's documentation.

Both *genecounting* and *hap* are able to handle SNPs and multiallelic markers, with the former be flexible enough to include features such as X-linked data and the later being able to handle large number of SNPs. But they are unable to recode allele labels automatically, so functions *gc.em* and *hap.em* are in *haplo.em* format and used by a modified function *hap.score* in association testing.

It is notable that multilocus data are handled differently from that in **hwde** and elegant definitions of basic genetic data can be found in **genetics** package.

Incidentally, I found my C mixed-radixed sorting routine as in Zhao and Sham (2003) is much faster than R's internal function.

With exceptions such as function *pfc* which is very computer-intensive, most functions in the package can easily be adapted for analysis of large datasets involving either SNPs or multiallelic markers. Some are utility functions, e.g. *muvar* and *whscore*, which will be part of the other analysis routines in the future.

For users, all functions have unified format. For developers, it is able to incorporate their C/C++ programs more easily and avoid repetitive work such as preparing own routines for matrix algebra and linear models. Further advantage can be taken from packages in **Bioconductor**, which are designed and written to deal with large number of genes.

3 Examples

Examples can be found from most function documentations. You can also try several simple examples via *demo*:

```
library(gap)
demo(gap)
```

4 Known bugs

Unaware of any bug. However, better memory management is expected.

5 Bibliographic note

The main references are Chow (1960), Guo and Thompson (1992), Williams et al. (1992), Gholamic and Thomas (1994), Risch and Merikangas (1996), Spielman and Ewens (1996), Risch and Merikangas (1997), Miller (1997), Sham (1997), Sham (1998), Devlin and Roeder (1999), Zhao et al. (1999), Guo and Lange (2000), Hirotsu et al. (2001), Zhao et al. (2002), Zaykin et al. (2002), Zhao (2004), Wacholder et al. (2004), Wang (2005), Skol et al. (2006), Wakefield (2007).

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