An Integrated Genetic Analysis Package Using R

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1 Introduction

This package was designed 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 in R, e.g. **genetics**, **hwde**, **haplo.score**, 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.

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

2 Implementation

The following, extracted from the package INDEX, shows the data and functions currently available.

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
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

familiality

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 equlibrium test for

multiallelic marker

hwe.hardy Hardy-Weinberg equlibrium 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

print.hap.score Print a hap.score object s2k Statistics for 2 by K table

snca A study of Parkinson's disease and SNCA makers

tbyt LD statistics for two SNPs

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().

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, an advantage over algorithms in **haplo.score**. But the latter is able to recode allele labels automatically, so functions gc.em and hap.em are in **haplo.score**'s 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 multial-lelic 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

whscore

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), 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).

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