

## ***adegenet* 1.3-1: new tools for the analysis of genome-wide SNP data**

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### **ABSTRACT**

**Summary:** While the R software is becoming a standard for the analysis of genetic data, classical population genetics tools are being challenged by the increasing availability of genomic sequences. Dedicated tools are needed for harnessing the large amount of information generated by next-generation sequencing technologies. We introduce new tools implemented in the *adegenet* 1.3-1 package for handling and analyzing genome-wide single nucleotide polymorphism (SNP) data. Using a bit-level coding scheme for SNP data and parallelized computation, *adegenet* enables the analysis of large genome-wide SNPs datasets using standard personal computers.

**Availability:** *adegenet* 1.3-1 is available from CRAN: <http://cran.r-project.org/web/packages/adegenet/>. Information and support including a dedicated forum of discussion can be found on the *adegenet* website: <http://adegenet.r-forge.r-project.org/>. *adegenet* is released with a manual and four tutorials totalling over 300 pages of documentation, and distributed under the GNU General Public Licence (≥2).

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### **1 INTRODUCTION**

The free software R (R Development Core Team, 2011) is becoming a standard for the analysis of genetic data, offering a wealth of packages dedicated to population genetics (Jombart, 2008; Paradis, 2010), phylogenetics (Paradis *et al.*, 2004; Schliep, 2011) or genome-wide association studies (Aulchenko *et al.*, 2007; Clayton and Leung, 2007). Until recently, classical genetic marker data such as microsatellites could be analyzed using standard tools and personal computers. However, the increasing availability of genomic sequence data has challenged both the tools and the resources needed to carry such analyses. While some specific packages have been developed for human association studies (Aulchenko *et al.*, 2007; Clayton and Leung, 2007), more general tools for the analysis of the genetic structure of biological populations are needed. In this article, we introduce new tools implemented in the R package *adegenet* (Jombart, 2008) which

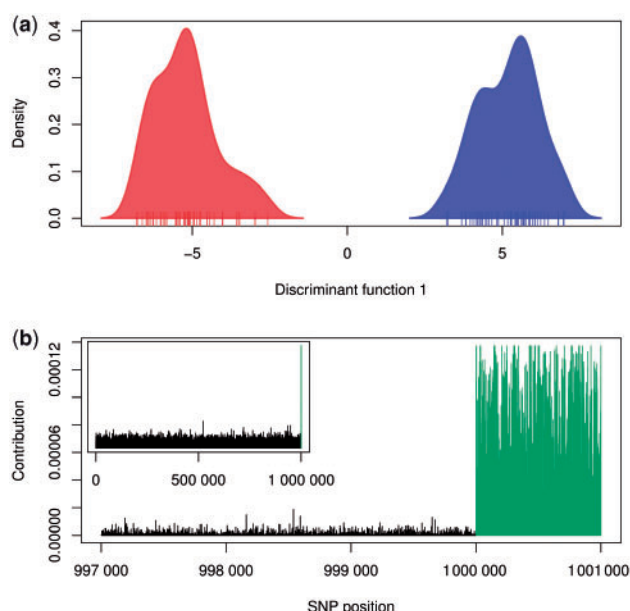
allow large genomic datasets (e.g. hundreds of individuals typed for hundreds of thousands SNPs) to be analyzed using standard personal computers. As an illustration, we show how a new implementation of the discriminant analysis of principal components (DAPC) (Jombart *et al.*, 2010) can be used to identify structuring alleles from genomic data with minimum computing resources.

### **2 DESCRIPTION**

The sheer size of genomic sequence data often precludes their analysis using standard personal computers. While studies focusing on genetic diversity can reduce the size of the analyzed datasets by considering biallelic SNPs only, the subsequent amount of data often remains considerable and can require prohibitive amounts of random access memory (RAM). To address this issue, we implemented a new data representation which codes each biallelic SNP using a single bit. While such coding is not readily possible in R, the new class *genlight* internally codes chunks of 8 SNPs using a single byte, resulting in drastic compression of the data. For instance, 50 individuals genotyped for 1 000 000 SNPs classically coded as characters would require ~380 MB of RAM, as opposed to 6 MB using *genlight* objects. This new coding scheme is also about eight times more compact than other available classes for representing SNP data such as *DNAbin* (Paradis *et al.*, 2004) or *snp.matrix* (Clayton and Leung, 2007). A further advantage of *genlight* is the ability to accommodate any ploidy in the data, even allowing for the ploidy to vary across individuals. The features of the class *genlight* are fully documented in a tutorial accessible from R by typing `vignette("adegenet-genomics")`.

While the bit-level coding of SNP data is undoubtedly memory efficient, it also makes the internal structure of the objects far more complex. Considerable efforts have been made to simplify the handling and analysis of *genlight* objects, whose manipulation is very close to matrices of individual allele frequencies. The entire *genlight* class has been replicated in C, which allowed for optimizing recurrent operations such as conversions from and to integers. Dedicated functions ('accessors') facilitate the access and modification of information while preventing the user from interacting directly with the complex internal structure of the objects. As a result, *genlight* objects act as 'black boxes' which resemble matrices of individual allele frequencies, albeit storing the information more efficiently. Basic functions such as mean and variance of SNP frequencies have also been implemented in order to facilitate the development of future dedicated tools.

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**Fig. 1.** DAPC of simulated data (see text). (a) Density of individual scores on the first discriminant function, with groups represented in red and blue. (b) SNP contribution to the separation of the groups; the last (structured) 1000 SNPs are coloured in green; the last 4000 SNPs are represented in the main plot, while the figure corresponding to all SNPs is shown in inset.

Beyond the need for efficient data storage, the analysis of genome-wide SNP data also requires significant computing power. Fortunately, most computers now possess processors with multiple cores, which can be used to partition important tasks into several smaller operations executed simultaneously by the different cores. This approach can lead to appreciable reductions in computational time and is most useful for analyzing large datasets. By default, most procedures implemented for *genlight* objects achieve parallelization using the package *multicore* (currently available on linux and MacOSX systems), although this can be disabled by the user. For instance, the new implementations of PCA (function *glPca*) and DAPC (*dapc*, see example) by default use compiled C code and parallelized computations, while never requiring more than two genomes to be represented as integers at a time. In some cases, this approach turns out to be even faster than other classical implementations of PCA (Supplementary Table S1).

Data interoperability can be a critical issue when large datasets are considered. Therefore, we made sure that genome-wide SNP data could be imported from standard formats into *genlight* objects as simply as possible. First, *genlight* objects can be created from lists or matrices of individual allele frequencies. Data can also be imported from the widely used software PLINK (Purcell *et al.*, 2007), which has defined a standard format for storing diploid SNP data. Alternatively, data can also be imported from *adegenet*'s own format (*.snp* files), which can accommodate any degree of ploidy and can store any meta-information such as individual group membership or positions of the SNPs. Finally, SNPs can also be directly extracted from aligned DNA sequences stored as FASTA files. Importantly, all these functions allow for processing the data by chunks of a few individuals, which allows for minimizing the RAM required for reading the data in.

### 3 EXAMPLE

We illustrate how a new implementation of DAPC for *genlight* objects can be used to identify structuring alleles from genome-wide SNP data. After loading the package, we simulate 1 001 000 SNPs for two groups of 50 individuals using *glSim*. The first 1 000 000 SNPs have similar distributions for both groups, whereas these distributions differ in the last 1000 SNPs.

```
> library(adegenet)
> x <- glSim(n.ind=100, n.snp.nonstruc=1e6,
             n.snp.struc=1e3)
```

We then apply DAPC to these data, choosing to retain 20 principal components in the prior dimension-reduction step.

```
> dapc1 <- dapc(x, n.pca=20)
```

Despite defavourable noise/signal ratio, DAPC discriminates very neatly the two groups of individuals (Fig. 1a). Interestingly, it also clearly identifies the structuring SNPs (Fig. 1b). Despite its simplicity, this example suggests that DAPC could be a useful tool for identifying structuring alleles from genome-wide SNP data.

### 4 CONCLUSION

*adegenet 1.3-1* provides new tools for the analysis of genome-wide SNP data using standard personal computers. As the availability of genomic data increases faster than computing resources, efficient data representation and parallel computation represent viable alternatives to the mere increase of raw computing power. As such, we hope that the new class *genlight* and the associated tools will make a significant contribution to taking population genetics studies into the genomic era and encourage the development of new dedicated methods.

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