

Overview of GGtools for expression genetics

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1 Introduction

The *GGtools* package contains infrastructure and demonstration data for joint analysis of transcriptome and genome through combination of DNA expression microarray and high-density SNP genotyping data. For Bioconductor 2.2 we adopted a representation of genotypes due to Clayton (in package *snpMatrix*) allowing reasonably convenient storage and manipulation of 4 megaSNP phase II HapMap genotypes on all the CEPH CEU samples. This contrasts with the previous version of *GGtools* which was limited to 550 kiloSNP and 58 CEU founders.

To give an immediate taste of the capabilities, we attach the package and load some test data.

```
> library(GGtools)
> data(hmceuB36.2021)
> hmceuB36.2021
```

```
snp.matrix-based genotype set:
number of samples: 90
number of chromosomes present: 2
annotation: illuminaHumanv1.db
Expression data dims: 47293 x 90
Phenodata: An object of class "AnnotatedDataFrame"
  sampleNames: NA06985, NA06991, ..., NA12892 (90 total)
  varLabels and varMetadata description:
    famid: hapmap family id
    persid: hapmap person id
    ...: ...
    male: logical TRUE if male
    (7 total)
```

Expression data are recoverable in a familiar way:

```
> exprs(hmceuB36.2021)[1:5, 1:5]
```

	NA06985	NA06991	NA06993	NA06994	NA07000
GI_10047089-S	5.983962	5.939529	5.912270	5.891347	5.906675
GI_10047091-S	6.544493	6.286516	6.244446	6.277397	6.330893
GI_10047093-S	9.905235	10.353804	10.380972	9.889223	10.155686
GI_10047099-S	7.993935	7.593970	8.261215	6.598430	6.728085
GI_10047103-S	11.882265	12.204753	12.249708	11.798415	12.015252

Genotype data have more complex representation.

```
> smList(hmceuB36.2021)
```

```
$`20`
```

```
A snp.matrix with 90 rows and 119921 columns
```

```
Row names: NA06985 ... NA12892
```

```
Col names: rs4814683 ... rs6090120
```

```
$`21`
```

```
A snp.matrix with 90 rows and 50165 columns
```

```
Row names: NA06985 ... NA12892
```

```
Col names: rs885550 ... rs10483083
```

```
> class(smList(hmceuB36.2021)[["20"]])
```

```
[1] "snp.matrix"
```

This shows that we use a named list to hold items of the *snp.matrix* class from *snpMatrix*.

It will generally be unnecessary to probe to this level, but it is instructive to check the underlying representation:

```
> schunk = smList(hmceuB36.2021)[["20"]]
```

```
> schunk@.Data[1:4, 1:4]
```

	rs4814683	rs6076506	rs6139074	rs1418258
NA06985	03	03	03	03
NA06991	02	03	02	02
NA06993	01	03	01	01
NA06994	01	03	01	01

The leading zeroes show that a raw byte representation is used. We can convert to allele codes as follows:

```
> as(schunk[1:4, 1:4], "character")
```

```

          rs4814683 rs6076506 rs6139074 rs1418258
NA06985 "B/B"      "B/B"      "B/B"      "B/B"
NA06991 "A/B"      "B/B"      "A/B"      "A/B"
NA06993 "A/A"      "B/B"      "A/A"      "A/A"
NA06994 "A/A"      "B/B"      "A/A"      "A/A"

```

The primary method of interest is the genome-wide association study, here applied with expression as the phenotype. Here we execute a founders-only analysis, adjusting for gender, confining attention to chromosome 20:

```

> pd = pData(hmceuB36.2021)
> hmFou = hmceuB36.2021[, which(pd$mothid == 0 & pd$fathid == 0)]
> f1 = gwSnpTests(genesym("CPNE1") ~ male, hmFou, chrnum(20))

```

2 Conversion to nucleotide codes

This is currently somewhat cumbersome. Suppose we want to know the specific nucleotide assignments for a given genotype call. For example, rs4814683 for subject NA06985.

```

> schunk["NA06985", "rs4814683"]

```

```

Autosomal snp(s):
[1] "B/B"

```

We need to know a) that the A/B tokens map in lexical order to the nucleotides (A will be the alphabetically first nucleotide for the diallelic call).

Using the `SNPlocs.Hsapiens.dbSNP.20071016` package, we can get the nucleotides:

```

> library(SNPlocs.Hsapiens.dbSNP.20071016)
> s20 = getSNPlocs("chr20")
> s20[s20[, 1] == 4814683, ]

```

```

RefSNP_id alleles_as_ambig loc
4 4814683 K 9795

```

Now we need to translate the IUPAC code to the nucleotides:

```

> library(Biostrings)
> IUPAC_CODE_MAP

```

```

      A      C      G      T      M      R      W      S      Y      K      V
"A"    "C"    "G"    "T"    "AC"    "AG"    "AT"    "CG"    "CT"    "GT"    "ACG"
      H      D      B      N
"ACT"  "AGT"  "CGT"  "ACGT"

```