

The biomaRt user's guide

Steffen Durinck*, Wolfgang Huber†

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*durincks@gene.com

†huber@ebi.ac.uk

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1 Introduction

In recent years a wealth of biological data has become available in public data repositories. Easy access to these valuable data resources and firm integration with data analysis is needed for comprehensive bioinformatics data analysis. The *biomaRt* package, provides an interface to a growing

collection of databases implementing the BioMart software suite (<http://www.biomart.org>). The package enables retrieval of large amounts of data in a uniform way without the need to know the underlying database schemas or write complex SQL queries. Examples of BioMart databases are Ensembl, Uniprot and HapMap. These major databases give biomaRt users direct access to a diverse set of data and enable a wide range of powerful online queries from R.

2 Selecting a BioMart database and dataset

Every analysis with *biomaRt* starts with selecting a BioMart database to use. A first step is to check which BioMart web services are available. The function `listMarts` will display all available BioMart web services

```
> library("biomaRt")
> listMarts()

      biomart          version
1 ENSEMBL_MART_ENSEMBL      Ensembl Genes 82
2   ENSEMBL_MART_SNP      Ensembl Variation 82
3 ENSEMBL_MART_FUNCGEN      Ensembl Regulation 82
4   ENSEMBL_MART_VEGA          Vega 62
5           pride          PRIDE (EBI UK)
```

Note: if the function `useMart` runs into proxy problems you should set your proxy first before calling any biomaRt functions. You can do this using the `Sys.putenv` command:

```
Sys.putenv("http\proxy" = "http://my.proxy.org:9999")
```

Some users have reported that the workaround above does not work, in this case an alternative proxy solution below can be tried:

```
options(RCurlOptions = list(proxy="uscache.kcc.com:80", proxyuserpwd="-----:-----"))
```

The `useMart` function can now be used to connect to a specified BioMart database, this must be a valid name given by `listMarts`. In the next example we choose to query the Ensembl BioMart database.

```
> ensembl=useMart("ensembl")
```

BioMart databases can contain several datasets, for Ensembl every species is a different dataset. In a next step we look at which datasets are available in the selected BioMart by using the function `listDatasets`.

> listDatasets(ensembl)

| | dataset | description | version |
|----|--------------------------------|---|-----------------|
| 1 | oanatinus_gene_ensembl | Ornithorhynchus anatinus genes (OANA5) | OANA5 |
| 2 | cporcellus_gene_ensembl | Cavia porcellus genes (cavPor3) | cavPor3 |
| 3 | gaculeatus_gene_ensembl | Gasterosteus aculeatus genes (BROADS1) | BROADS1 |
| 4 | lafricana_gene_ensembl | Loxodonta africana genes (loxAfr3) | loxAfr3 |
| 5 | itridecemlineatus_gene_ensembl | Ictidomys tridecemlineatus genes (spetri2) | spetri2 |
| 6 | choffmanni_gene_ensembl | Choloepus hoffmanni genes (choHof1) | choHof1 |
| 7 | csavignyi_gene_ensembl | Ciona savignyi genes (CSAV2.0) | CSAV2.0 |
| 8 | fcatus_gene_ensembl | Felis catus genes (Felis_catus_6.2) | Felis_catus_6.2 |
| 9 | rnorvegicus_gene_ensembl | Rattus norvegicus genes (Rnor_6.0) | Rnor_6.0 |
| 10 | psinensis_gene_ensembl | Pelodiscus sinensis genes (PelSin_1.0) | PelSin_1.0 |
| 11 | cjacchus_gene_ensembl | Callithrix jacchus genes (C_jacchus3.2.1) | C_jacchus3.2.1 |
| 12 | ttruncatus_gene_ensembl | Tursiops truncatus genes (turTru1) | turTru1 |
| 13 | scerevisiae_gene_ensembl | Saccharomyces cerevisiae genes (R64-1-1) | R64-1-1 |
| 14 | celegans_gene_ensembl | Caenorhabditis elegans genes (WBcel235) | WBcel235 |
| 15 | csabaeus_gene_ensembl | Chlorocebus sabaues genes (ChlSab1.1) | ChlSab1.1 |
| 16 | oniloticus_gene_ensembl | Oreochromis niloticus genes (Orenil1.0) | Orenil1.0 |
| 17 | trubripes_gene_ensembl | Takifugu rubripes genes (FUGU4.0) | FUGU4.0 |
| 18 | amexicanus_gene_ensembl | Astyanax mexicanus genes (AstMex102) | AstMex102 |
| 19 | pmarinus_gene_ensembl | Petromyzon marinus genes (Pmarinus_7.0) | Pmarinus_7.0 |
| 20 | eeuropaeus_gene_ensembl | Erinaceus europaeus genes (eriEur1) | eriEur1 |
| 21 | falbicollis_gene_ensembl | Ficedula albicollis genes (FicAlb_1.4) | FicAlb_1.4 |
| 22 | ptroglodytes_gene_ensembl | Pan troglodytes genes (CHIMP2.1.4) | CHIMP2.1.4 |
| 23 | etelfairi_gene_ensembl | Echinops telfairi genes (TENREC) | TENREC |
| 24 | cintestinalis_gene_ensembl | Ciona intestinalis genes (KH) | KH |
| 25 | nleucogenys_gene_ensembl | Nomascus leucogenys genes (Nleu1.0) | Nleu1.0 |
| 26 | sscrofa_gene_ensembl | Sus scrofa genes (Sscrofa10.2) | Sscrofa10.2 |
| 27 | ocuniculus_gene_ensembl | Oryctolagus cuniculus genes (OryCun2.0) | OryCun2.0 |
| 28 | dnovemcinctus_gene_ensembl | Dasyops novemcinctus genes (Dasnov3.0) | Dasnov3.0 |
| 29 | pcapensis_gene_ensembl | Procavia capensis genes (proCap1) | proCap1 |
| 30 | tguttata_gene_ensembl | Taeniopygia guttata genes (taeGut3.2.4) | taeGut3.2.4 |
| 31 | mlucifugus_gene_ensembl | Myotis lucifugus genes (myoLuc2) | myoLuc2 |
| 32 | hsapiens_gene_ensembl | Homo sapiens genes (GRCh38.p3) | GRCh38.p3 |
| 33 | pfiformosa_gene_ensembl | Poecilia formosa genes (PoeFor_5.1.2) | PoeFor_5.1.2 |
| 34 | mfuro_gene_ensembl | Mustela putorius furo genes (MusPutFur1.0) | MusPutFur1.0 |
| 35 | tbelangeri_gene_ensembl | Tupaia belangeri genes (tupBel1) | tupBel1 |
| 36 | ggallus_gene_ensembl | Gallus gallus genes (Galgal4) | Galgal4 |
| 37 | xtropicalis_gene_ensembl | Xenopus tropicalis genes (JGI4.2) | JGI4.2 |
| 38 | ecaballus_gene_ensembl | Equus caballus genes (EquCab2) | EquCab2 |
| 39 | pabelii_gene_ensembl | Pongo abelii genes (PPYG2) | PPYG2 |
| 40 | xmaculatus_gene_ensembl | Xiphophorus maculatus genes (Xipmac4.4.2) | Xipmac4.4.2 |
| 41 | drerio_gene_ensembl | Danio rerio genes (GRCz10) | GRCz10 |
| 42 | lchalumnae_gene_ensembl | Latimeria chalumnae genes (LatCha1) | LatCha1 |
| 43 | tnigroviridis_gene_ensembl | Tetraodon nigroviridis genes (TETRAODON8.0) | TETRAODON8.0 |
| 44 | amelanoleuca_gene_ensembl | Ailuropoda melanoleuca genes (ailMel1) | ailMel1 |
| 45 | mmulatta_gene_ensembl | Macaca mulatta genes (MMUL_1) | MMUL_1 |
| 46 | pvampyrus_gene_ensembl | Pteropus vampyrus genes (pteVam1) | pteVam1 |
| 47 | panubis_gene_ensembl | Papio anubis genes (PapAnu2.0) | PapAnu2.0 |
| 48 | mdomestica_gene_ensembl | Monodelphis domestica genes (monDom5) | monDom5 |
| 49 | acarolinensis_gene_ensembl | Anolis carolinensis genes (AnoCar2.0) | AnoCar2.0 |
| 50 | vpacos_gene_ensembl | Vicugna pacos genes (vicPac1) | vicPac1 |
| 51 | tsyrichta_gene_ensembl | Tarsius syrichta genes (tarSyr1) | tarSyr1 |
| 52 | ogarnettii_gene_ensembl | Otolemur garnettii genes (OtoGar3) | OtoGar3 |
| 53 | dmelanogaster_gene_ensembl | Drosophila melanogaster genes (BDGP6) | BDGP6 |
| 54 | mmurinus_gene_ensembl | Microcebus murinus genes (micMur1) | micMur1 |

| | | | |
|----|-----------------------------|---|--------------|
| 55 | loculatus_gene_ensembl | Lepisosteus oculatus genes (LepOcu1) | LepOcu1 |
| 56 | olatipes_gene_ensembl | Oryzias latipes genes (HdrR) | HdrR |
| 57 | ggorilla_gene_ensembl | Gorilla gorilla genes (gorGor3.1) | gorGor3.1 |
| 58 | oprinceps_gene_ensembl | Ochotona princeps genes (OchPri2.0) | OchPri2.0 |
| 59 | dordii_gene_ensembl | Dipodomys ordii genes (dipOrd1) | dipOrd1 |
| 60 | oaries_gene_ensembl | Ovis aries genes (Oar_v3.1) | Oar_v3.1 |
| 61 | mmusculus_gene_ensembl | Mus musculus genes (GRCm38.p4) | GRCm38.p4 |
| 62 | mgallopavo_gene_ensembl | Meleagris gallopavo genes (UMD2) | UMD2 |
| 63 | gmorhua_gene_ensembl | Gadus morhua genes (gadMor1) | gadMor1 |
| 64 | aplatyrhynchos_gene_ensembl | Anas platyrhynchos genes (BGI_duck_1.0) | BGI_duck_1.0 |
| 65 | saraneus_gene_ensembl | Sorex araneus genes (sorAra1) | sorAra1 |
| 66 | sharrisii_gene_ensembl | Sarcophilus harrisii genes (DEVIL7.0) | DEVIL7.0 |
| 67 | meugenii_gene_ensembl | Macropus eugenii genes (Meug_1.0) | Meug_1.0 |
| 68 | btaurus_gene_ensembl | Bos taurus genes (UMD3.1) | UMD3.1 |
| 69 | cfamiliaris_gene_ensembl | Canis familiaris genes (CanFam3.1) | CanFam3.1 |

To select a dataset we can update the `Mart` object using the function `useDataset`. In the example below we choose to use the `hsapiens` dataset.

```
ensembl = useDataset("hsapiens_gene_ensembl", mart=ensembl)
```

Or alternatively if the dataset one wants to use is known in advance, we can select a BioMart database and dataset in one step by:

```
> ensembl = useMart("ensembl", dataset="hsapiens_gene_ensembl")
```

3 How to build a biomaRt query

The `getBM` function has three arguments that need to be introduced: filters, attributes and values. *Filters* define a restriction on the query. For example you want to restrict the output to all genes located on the human X chromosome then the filter `chromosome_name` can be used with value 'X'. The `listFilters` function shows you all available filters in the selected dataset.

```
> filters = listFilters(ensembl)
> filters[1:5,]
```

| | name | description |
|---|-----------------|-----------------|
| 1 | chromosome_name | Chromosome name |
| 2 | start | Gene Start (bp) |
| 3 | end | Gene End (bp) |
| 4 | band_start | Band Start |
| 5 | band_end | Band End |

Attributes define the values we are interested in to retrieve. For example we want to retrieve the gene symbols or chromosomal coordinates. The `listAttributes` function displays all available attributes in the selected dataset.

```
> attributes = listAttributes(ensembl)
> attributes[1:5,]
```

| | name | description |
|---|-----------------------|-----------------------|
| 1 | ensembl_gene_id | Ensembl Gene ID |
| 2 | ensembl_transcript_id | Ensembl Transcript ID |
| 3 | ensembl_peptide_id | Ensembl Protein ID |
| 4 | ensembl_exon_id | Ensembl Exon ID |
| 5 | description | Description |

The `getBM` function is the main query function in `biomaRt`. It has four main arguments:

- `attributes`: is a vector of attributes that one wants to retrieve (= the output of the query).
- `filters`: is a vector of filters that one will use as input to the query.
- `values`: a vector of values for the filters. In case multiple filters are in use, the `values` argument requires a list of values where each position in the list corresponds to the position of the filters in the `filters` argument (see examples below).
- `mart`: is an object of class `Mart`, which is created by the `useMart` function.

Note: for some frequently used queries to Ensembl, wrapper functions are available: `getGene` and `getSequence`. These functions call the `getBM` function with hard coded filter and attribute names.

Now that we selected a BioMart database and dataset, and know about attributes, filters, and the values for filters; we can build a `biomaRt` query. Let's make an easy query for the following problem: We have a list of Affymetrix identifiers from the `u133plus2` platform and we want to retrieve the corresponding EntrezGene identifiers using the Ensembl mappings.

The `u133plus2` platform will be the filter for this query and as values for this filter we use our list of Affymetrix identifiers. As output (attributes) for

the query we want to retrieve the EntrezGene and u133plus2 identifiers so we get a mapping of these two identifiers as a result. The exact names that we will have to use to specify the attributes and filters can be retrieved with the `listAttributes` and `listFilters` function respectively. Let's now run the query:

```
> affyids=c("202763_at","209310_s_at","207500_at")
> getBM(attributes=c('affy_hg_u133_plus_2', 'entrezgene'), filters = 'affy_hg_u133_plus_2', values = affyids, mart =

  affy_hg_u133_plus_2  entrezgene
1      209310_s_at      837
2      207500_at      838
3      202763_at      836
```

4 Examples of biomaRt queries

In the sections below a variety of example queries are described. Every example is written as a task, and we have to come up with a biomaRt solution to the problem.

4.1 Task 1: Annotate a set of Affymetrix identifiers with HUGO symbol and chromosomal locations of corresponding genes

We have a list of Affymetrix hgu133plus2 identifiers and we would like to retrieve the HUGO gene symbols, chromosome names, start and end positions and the bands of the corresponding genes. The `listAttributes` and the `listFilters` functions give us an overview of the available attributes and filters and we look in those lists to find the corresponding attribute and filter names we need. For this query we'll need the following attributes: `hgnc_symbol`, `chromosome_name`, `start_position`, `end_position`, `band` and `affy_hg_u133_plus_2` (as we want these in the output to provide a mapping with our original Affymetrix input identifiers. There is one filter in this query which is the `affy_hg_u133_plus_2` filter as we use a list of Affymetrix identifiers as input. Putting this all together in the `getBM` and performing the query gives:

```
> affyids=c("202763_at","209310_s_at","207500_at")
> getBM(attributes=c('affy_hg_u133_plus_2', 'hgnc_symbol', 'chromosome_name', 'start_position', 'end_position', 'band'),
+ filters = 'affy_hg_u133_plus_2', values = affyids, mart = ensembl)

  affy_hg_u133_plus_2 hgnc_symbol chromosome_name start_position end_position  band
1      209310_s_at      CASP4             11      104813593  104840163 q22.3
2      207500_at      CASP5             11      104864962  104893895 q22.3
3      202763_at      CASP3              4      185548850  185570663 q35.1
```

4.2 Task 2: Annotate a set of EntrezGene identifiers with GO annotation

In this task we start out with a list of EntrezGene identifiers and we want to retrieve GO identifiers related to biological processes that are associated with these entrezgene identifiers. Again we look at the output of `listAttributes` and `listFilters` to find the filter and attributes we need. Then we construct the following query:

```
> entrez=c("673","837")
> goids = getBM(attributes=c('entrezgene','go_id'), filters='entrezgene', values=entrez, mart=ensembl)
> head(goids)

  entrezgene   go_id
1         673 GO:0000186
2         673 GO:0006468
3         673 GO:0006916
4         673 GO:0007264
5         673 GO:0007268
```

4.3 Task 3: Retrieve all HUGO gene symbols of genes that are located on chromosomes 17,20 or Y , and are associated with one the following GO terms: "GO:0051330","GO:0000080","GO:0000114","GO:0000082" (here we'll use more than one filter)

The `getBM` function enables you to use more than one filter. In this case the filter argument should be a vector with the filter names. The values should be a list, where the first element of the list corresponds to the first filter and the second list element to the second filter and so on. The elements of this list are vectors containing the possible values for the corresponding filters.

```
go=c("GO:0051330","GO:0000080","GO:0000114")
chrom=c(17,20,"Y")
getBM(attributes= "hgnc_symbol",
       filters=c("go_id","chromosome_name"),
       values=list(go,chrom), mart=ensembl)

  hgnc_symbol
1         E2F1
```

4.4 Task 4: Annotate set of identifiers with INTERPRO protein domain identifiers

In this example we want to annotate the following two RefSeq identifiers: NM_005359 and NM_000546 with INTERPRO protein domain identifiers and a description of the protein domains.

```

> refseqids = c("NM_005359", "NM_000546")
> ipro = getBM(attributes=c("refseq_dna", "interpro", "interpro_description"), filters=

ipro
  refseq_dna  interpro          interpro_description
1 NM_000546  IPR002117          p53 tumor antigen
2 NM_000546  IPR010991          p53, tetramerisation
3 NM_000546  IPR011615          p53, DNA-binding
4 NM_000546  IPR013872  p53 transactivation domain (TAD)
5 NM_000546  IPR000694          Proline-rich region
6 NM_005359  IPR001132          MAD homology 2, Dwarfing-type
7 NM_005359  IPR003619          MAD homology 1, Dwarfing-type
8 NM_005359  IPR013019          MAD homology, MH1

```

4.5 Task 5: Select all Affymetrix identifiers on the hgu133plus2 chip and Ensembl gene identifiers for genes located on chromosome 16 between basepair 1100000 and 1250000.

In this example we will again use multiple filters: chromosome_name, start, and end as we filter on these three conditions. Note that when a chromosome name, a start position and an end position are jointly used as filters, the BioMart webservice interprets this as return everything from the given chromosome between the given start and end positions.

```

> getBM(c('affy_hg_u133_plus_2', 'ensembl_gene_id'), filters = c('chromosome_name', 'start', 'end'),
+ values=list(16, 1100000, 1250000), mart=ensembl)

```

```

  affy_hg_u133_plus_2  ensembl_gene_id
1                    ENSG00000260702
2          215502_at  ENSG00000260532
3                    ENSG00000273551
4          205845_at  ENSG00000196557
5                    ENSG00000196557
6                    ENSG00000260403
7                    ENSG00000259910
8                    ENSG00000261294
9          220339_s_at  ENSG00000116176
10                   ENSG00000277010
11          217023_x_at  ENSG00000197253
12          210084_x_at  ENSG00000197253
13          215382_x_at  ENSG00000197253
14          216474_x_at  ENSG00000197253
15          207134_x_at  ENSG00000197253
16          205683_x_at  ENSG00000197253
17          217023_x_at  ENSG00000172236
18          210084_x_at  ENSG00000172236
19          215382_x_at  ENSG00000172236
20          207741_x_at  ENSG00000172236
21          216474_x_at  ENSG00000172236
22          207134_x_at  ENSG00000172236
23          205683_x_at  ENSG00000172236

```

4.6 Task 6: Retrieve all entrezgene identifiers and HUGO gene symbols of genes which have a "MAP kinase activity" GO term associated with it.

The GO identifier for MAP kinase activity is GO:0004707. In our query we will use go as filter and entrezgene and hgnc_symbol as attributes. Here's the query:

```
> getBM(c('entrezgene','hgnc_symbol'), filters='go', values='GO:0004707', mart=ensembl)
```

| | entrezgene | hgnc_symbol |
|---|------------|-------------|
| 1 | 5601 | MAPK9 |
| 2 | 225689 | MAPK15 |
| 3 | 5599 | MAPK8 |
| 4 | 5594 | MAPK1 |
| 5 | 6300 | MAPK12 |

4.7 Task 7: Given a set of EntrezGene identifiers, retrieve 100bp upstream promoter sequences

All sequence related queries to Ensembl are available through the `getSequence` wrapper function. `getBM` can also be used directly to retrieve sequences but this can get complicated so using `getSequence` is recommended. Sequences can be retrieved using the `getSequence` function either starting from chromosomal coordinates or identifiers. The chromosome name can be specified using the *chromosome* argument. The *start* and *end* arguments are used to specify *start* and *end* positions on the chromosome. The type of sequence returned can be specified by the *seqType* argument which takes the following values: 'cdna'; 'peptide' for protein sequences; '3utr' for 3' UTR sequences, '5utr' for 5' UTR sequences; 'gene_exon' for exon sequences only; 'transcript_exon' for transcript specific exonic sequences only; 'transcript_exon_intron' gives the full unspliced transcript, that is exons + introns; 'gene_exon_intron' gives the exons + introns of a gene; 'coding' gives the coding sequence only; 'coding_transcript_flank' gives the flanking region of the transcript including the UTRs, this must be accompanied with a given value for the upstream or downstream attribute; 'coding_gene_flank' gives the flanking region of the gene including the UTRs, this must be accompanied with a given value for the upstream or downstream attribute; 'transcript_flank' gives the flanking region of the transcript excluding the UTRs, this must be accompanied with a given value for the upstream or downstream attribute; 'gene_flank' gives the flanking region of the gene excluding the UTRs, this must be accompanied with a given value for the upstream or downstream attribute.

In MySQL mode the `getSequence` function is more limited and the sequence

that is returned is the 5' to 3'+ strand of the genomic sequence, given a chromosome, as start and an end position.

Task 4 requires us to retrieve 100bp upstream promoter sequences from a set of EntrzGene identifiers. The type argument in `getSequence` can be thought of as the filter in this query and uses the same input names given by `listFilters`. In our query we use `entrezgene` for the type argument. Next we have to specify which type of sequences we want to retrieve, here we are interested in the sequences of the promoter region, starting right next to the coding start of the gene. Setting the `seqType` to `coding_gene_flank` will give us what we need. The `upstream` argument is used to specify how many bp of upstream sequence we want to retrieve, here we'll retrieve a rather short sequence of 100bp. Putting this all together in `getSequence` gives:

```
> entrez=c("673", "7157", "837")
> getSequence(id = entrez, type="entrezgene", seqType="coding_gene_flank", upstream=100, mart=ensembl)
```

4.8 Task 8: Retrieve all 5' UTR sequences of all genes that are located on chromosome 3 between the positions 185514033 and 185535839

As described in the previous task `getSequence` can also use chromosomal coordinates to retrieve sequences of all genes that lie in the given region. We also have to specify which type of identifier we want to retrieve together with the sequences, here we choose for `entrezgene` identifiers.

```
> utr5 = getSequence(chromosome=3, start=185514033, end=185535839,
+                   type="entrezgene", seqType="5utr", mart=ensembl)
> utr5
```

```
          V1          V2
.....GAAGCGGTGGC .... 1981
```

4.9 Task 9: Retrieve protein sequences for a given list of EntrezGene identifiers

In this task the type argument specifies which type of identifiers we are using. To get an overview of other valid identifier types we refer to the `listFilters` function.

```
> protein = getSequence(id=c(100, 5728), type="entrezgene",
+                       seqType="peptide", mart=ensembl)
> protein
```

```

peptide      entrezgene
MAQTPAFDKPKVEL ... 100
MTAIIKEIVSRNKRR ... 5728

```

4.10 Task 10: Retrieve known SNPs located on the human chromosome 8 between positions 148350 and 148612

For this example we'll first have to connect to a different BioMart database, namely snp.

```
> snpmart = useMart("snp", dataset="hsapiens_snp")
```

The `listAttributes` and `listFilters` functions give us an overview of the available attributes and filters. From these we need: `refsnp_id`, `allele`, `chrom_start` and `chrom_strand` as attributes; and as filters we'll use: `chrom_start`, `chrom_end` and `chr_name`. Note that when a chromosome name, a start position and an end position are jointly used as filters, the BioMart webservice interprets this as return everything from the given chromosome between the given start and end positions. Putting our selected attributes and filters into `getBM` gives:

```
> getBM(c('refsnp_id', 'allele', 'chrom_start', 'chrom_strand'), filters = c('chr_name', 'chrom_start', 'chrom_end'), val
```

| | refsnp_id | allele | chrom_start | chrom_strand |
|----|------------|--------|-------------|--------------|
| 1 | rs1134195 | G/T | 148394 | -1 |
| 2 | rs4046274 | C/A | 148394 | 1 |
| 3 | rs4046275 | A/G | 148411 | 1 |
| 4 | rs13291 | C/T | 148462 | 1 |
| 5 | rs1134192 | G/A | 148462 | -1 |
| 6 | rs4046276 | C/T | 148462 | 1 |
| 7 | rs12019378 | T/G | 148471 | 1 |
| 8 | rs1134191 | C/T | 148499 | -1 |
| 9 | rs4046277 | G/A | 148499 | 1 |
| 10 | rs11136408 | G/A | 148525 | 1 |
| 11 | rs1134190 | C/T | 148533 | -1 |
| 12 | rs4046278 | G/A | 148533 | 1 |
| 13 | rs1134189 | G/A | 148535 | -1 |
| 14 | rs3965587 | C/T | 148535 | 1 |
| 15 | rs1134187 | G/A | 148539 | -1 |
| 16 | rs1134186 | T/C | 148569 | 1 |
| 17 | rs4378731 | G/A | 148601 | 1 |

4.11 Task 11: Given the human gene TP53, retrieve the human chromosomal location of this gene and also retrieve the chromosomal location and RefSeq id of it's homolog in mouse.

The `getLDS` (Get Linked Dataset) function provides functionality to link 2 BioMart datasets which each other and construct a query over the two

datasets. In Ensembl, linking two datasets translates to retrieving homology data across species. The usage of `getLDS` is very similar to `getBM`. The linked dataset is provided by a separate `Mart` object and one has to specify filters and attributes for the linked dataset. Filters can either be applied to both datasets or to one of the datasets. Use the `listFilters` and `listAttributes` functions on both `Mart` objects to find the filters and attributes for each dataset (species in Ensembl). The attributes and filters of the linked dataset can be specified with the `attributesL` and `filtersL` arguments. Entering all this information into `getLDS` gives:

```
human = useMart("ensembl", dataset = "hsapiens_gene_ensembl")
mouse = useMart("ensembl", dataset = "mmusculus_gene_ensembl")
getLDS(attributes = c("hgnc_symbol","chromosome_name", "start_position"),
        filters = "hgnc_symbol", values = "TP53",mart = human,
        attributesL = c("refseq_dna","chromosome_name","start_position"), martL = mouse)

      V1 V2      V3      V4 V5      V6
1 TP53 17 7512464 NM_011640 11 69396600
```

5 Using archived versions of Ensembl

It is possible to query archived versions of Ensembl through *biomaRt*. There are currently two ways to access archived versions.

5.1 Using the `archive=TRUE`

First we list the available Ensembl archives by using the `listMarts` function and setting the `archive` attribute to `TRUE`. Note that not all archives are available this way and it seems that recently this only gives access to few archives if you don't see the version of the archive you need please look at the 2nd way to access archives.

```
> listMarts(archive=TRUE)

      biomart      version
1      ensembl_mart_51      Ensembl 51
2      snp_mart_51      SNP 51
3      vega_mart_51      Vega 32
4      ensembl_mart_50      Ensembl 50
5      snp_mart_50      SNP 50
6      vega_mart_50      Vega 32
7      ensembl_mart_49      ENSEMBL GENES 49 (SANGER)
8      genomic_features_mart_49      Genomic Features
9      snp_mart_49      SNP
10     vega_mart_49      Vega
11     ensembl_mart_48      ENSEMBL GENES 48 (SANGER)
12     genomic_features_mart_48      Genomic Features
13     snp_mart_48      SNP
14     vega_mart_48      Vega
15     ensembl_mart_47      ENSEMBL GENES 47 (SANGER)
16     genomic_features_mart_47      Genomic Features
```

```

17          snp_mart_47          SNP
18          vega_mart_47          Vega
19  compara_mart_homology_47      Compara homology
20 compara_mart_multiple_ga_47    Compara multiple alignments
21 compara_mart_pairwise_ga_47    Compara pairwise alignments
22          ensembl_mart_46      ENSEMBL GENES 46 (SANGER)
23  genomic_features_mart_46      Genomic Features
24          snp_mart_46          SNP
25          vega_mart_46          Vega
26  compara_mart_homology_46      Compara homology
27 compara_mart_multiple_ga_46    Compara multiple alignments
28 compara_mart_pairwise_ga_46    Compara pairwise alignments
29          ensembl_mart_45      ENSEMBL GENES 45 (SANGER)
30          snp_mart_45          SNP
31          vega_mart_45          Vega
32  compara_mart_homology_45      Compara homology
33 compara_mart_multiple_ga_45    Compara multiple alignments
34 compara_mart_pairwise_ga_45    Compara pairwise alignments
35          ensembl_mart_44      ENSEMBL GENES 44 (SANGER)
36          snp_mart_44          SNP
37          vega_mart_44          Vega
38  compara_mart_homology_44      Compara homology
39 compara_mart_pairwise_ga_44    Compara pairwise alignments
40          ensembl_mart_43      ENSEMBL GENES 43 (SANGER)
41          snp_mart_43          SNP
42          vega_mart_43          Vega
43  compara_mart_homology_43      Compara homology
44 compara_mart_pairwise_ga_43    Compara pairwise alignments

```

Next we select the archive we want to use using the `useMart` function, again setting the `archive` attribute to `TRUE` and giving the full name of the BioMart e.g. `ensembl_mart_46`.

```
> ensembl = useMart("ensembl_mart_46", dataset="hsapiens_gene_ensembl", archive = TRUE)
```

If you don't know the dataset you want to use could first connect to the BioMart using `useMart` and then use the `listDatasets` function on this object. After you selected the BioMart database and dataset, queries can be performed in the same way as when using the current BioMart versions.

5.2 Accessing archives through specifying the archive host

Use the <http://www.ensembl.org> website and go down the bottom of the page. Click on 'view in Archive' and select the archive you need. Copy the url and use that url as shown below to connect to the specified BioMart database. The example below shows how to query Ensembl 54.

```
> listMarts(host='may2009.archive.ensembl.org')
> ensembl54=useMart(host='may2009.archive.ensembl.org', biomart='ENSEMBL_MART_ENSEMBL')
> ensembl54=useMart(host='may2009.archive.ensembl.org', biomart='ENSEMBL_MART_ENSEMBL', dataset='hsapiens_gene_ensembl')
```

6 Using a BioMart other than Ensembl

To demonstrate the use of the `biomaRt` package with non-Ensembl databases the next query is performed using the Wormbase BioMart (WormMart). We connect to Wormbase, select the gene dataset to use and have a look at the available attributes and filters. Then we use a list of gene names as filter and retrieve associated RNAi identifiers together with a description of the RNAi phenotype.

```
> wormbase=useMart("WS220",dataset="wormbase_gene")
> listFilters(wormbase)
> listAttributes(wormbase)
> getBM(attributes = c("public_name","rna_i","rna_i_phenotype_phenotype_label"),
+         filters="gene_name", values=c("unc-26","his-33"),
+         mart=wormbase)
>
```

| | public_name | rna_i | rna_i_phenotype_phenotype_label |
|---|-------------|----------------|-----------------------------------|
| 1 | his-33 | WBRNAi00082060 | GRO slow growth |
| 2 | his-33 | WBRNAi00082060 | postembryonic development variant |
| 3 | his-33 | WBRNAi00082060 | EMB embryonic lethal |
| 4 | his-33 | WBRNAi00082060 | LVL larval lethal |
| 5 | his-33 | WBRNAi00082060 | LVA larval arrest |
| 6 | his-33 | WBRNAi00082060 | accumulated cell corpses |

7 biomaRt helper functions

This section describes a set of `biomaRt` helper functions that can be used to export FASTA format sequences, retrieve values for certain filters and exploring the available filters and attributes in a more systematic manner.

7.1 exportFASTA

The data.frames obtained by the `getSequence` function can be exported to FASTA files using the `exportFASTA` function. One has to specify the data.frame to export and the filename using the `file` argument.

7.2 Finding out more information on filters

7.2.1 filterType

Boolean filters need a value `TRUE` or `FALSE` in `biomaRt`. Setting the value `TRUE` will include all information that fulfill the filter requirement. Setting `FALSE` will exclude the information that fulfills the filter requirement and will return all values that don't fulfill the filter. For most of the filters, their

name indicates if the type is a boolean or not and they will usually start with "with". However this is not a rule and to make sure you got the type right you can use the function `filterType` to investigate the type of the filter you want to use.

```
> filterType("with_affy_hg_u133_plus_2",ensembl)

[1] "boolean_list"
```

7.2.2 filterOptions

Some filters have a limited set of values that can be given to them. To know which values these are one can use the `filterOptions` function to retrieve the predetermined values of the respective filter.

```
> filterOptions("biotype",ensembl)

[1] "[3prime_overlapping_ncrna,antisense,IG_C_gene,IG_C_pseudogene,IG_D_gene,IG_J_gene,IG_J_p
```

If there are no predetermined values e.g. for the `entrezgene` filter, then `filterOptions` will return the type of filter it is. And most of the times the filter name or it's description will suggest what values one case use for the respective filter (e.g. `entrezgene` filter will work with `entrezgene` identifiers as values)

7.3 Attribute Pages

For large BioMart databases such as Ensembl, the number of attributes displayed by the `listAttributes` function can be very large. In BioMart databases, attributes are put together in pages, such as sequences, features, homologs for Ensembl. An overview of the attributes pages present in the respective BioMart dataset can be obtained with the `attributePages` function.

```
> pages = attributePages(ensembl)
> pages

[1] "feature_page" "structure"      "homologs"      "snp"           "snp_somatic"  "sequences"
```

To show us a smaller list of attributes which belong to a specific page, we can now specify this in the `listAttributes` function as follows:

```
> listAttributes(ensembl, page="feature_page")
```

| | name | description |
|----|---------------------------------|--|
| 1 | ensembl_gene_id | Ensembl Gene ID |
| 2 | ensembl_transcript_id | Ensembl Transcript ID |
| 3 | ensembl_peptide_id | Ensembl Protein ID |
| 4 | ensembl_exon_id | Ensembl Exon ID |
| 5 | description | Description |
| 6 | chromosome_name | Chromosome Name |
| 7 | start_position | Gene Start (bp) |
| 8 | end_position | Gene End (bp) |
| 9 | strand | Strand |
| 10 | band | Band |
| 11 | transcript_start | Transcript Start (bp) |
| 12 | transcript_end | Transcript End (bp) |
| 13 | transcription_start_site | Transcription Start Site (TSS) |
| 14 | transcript_length | Transcript length (including UTRs and CDS) |
| 15 | transcript_tsl | Transcript Support Level (TSL) |
| 16 | transcript_gencode_basic | GENCODE basic annotation |
| 17 | transcript_appris | APPRIS annotation |
| 18 | external_gene_name | Associated Gene Name |
| 19 | external_gene_source | Associated Gene Source |
| 20 | external_transcript_name | Associated Transcript Name |
| 21 | external_transcript_source_name | Associated Transcript Source |
| 22 | transcript_count | Transcript count |
| 23 | percentage_gc_content | % GC content |
| 24 | gene_biotype | Gene type |
| 25 | transcript_biotype | Transcript type |
| 26 | source | Source (gene) |
| 27 | transcript_source | Source (transcript) |
| 28 | status | Status (gene) |
| 29 | transcript_status | Status (transcript) |
| 30 | version | Version (gene) |
| 31 | transcript_version | Version (transcript) |
| 32 | phenotype_description | Phenotype description |
| 33 | source_name | Source name |
| 34 | study_external_id | Study External Reference |
| 35 | go_id | GO Term Accession |
| 36 | name_1006 | GO Term Name |
| 37 | definition_1006 | GO Term Definition |
| 38 | go_linkage_type | GO Term Evidence Code |
| 39 | namespace_1003 | GO domain |
| 40 | goslim_goa_accession | GOSlim GOA Accession(s) |
| 41 | goslim_goa_description | GOSlim GOA Description |
| 42 | arrayexpress | ArrayExpress |
| 43 | chembl | ChEMBL ID(s) |
| 44 | clone_based_ensembl_gene_name | Clone based Ensembl gene name |

| | | |
|----|-------------------------------------|---|
| 45 | clone_based_ensembl_transcript_name | Clone based Ensembl transcript name |
| 46 | clone_based_vega_gene_name | Clone based VEGA gene name |
| 47 | clone_based_vega_transcript_name | Clone based VEGA transcript name |
| 48 | ccds | CCDS ID |
| 49 | dbass3_id | Database of Aberrant 3' Splice Sites (DBASS3) IDs |
| 50 | dbass3_name | DBASS3 Gene Name |
| 51 | dbass5_id | Database of Aberrant 5' Splice Sites (DBASS5) IDs |
| 52 | dbass5_name | DBASS5 Gene Name |
| 53 | embl | EMBL (Genbank) ID |
| 54 | ens_hs_transcript | Ensembl Human Transcript IDs |
| 55 | ens_hs_translation | Ensembl Human Translation IDs |
| 56 | ens_lrg_gene | LRG to Ensembl link gene |
| 57 | ens_lrg_transcript | LRG to Ensembl link transcript |
| 58 | entrezgene | EntrezGene ID |
| 59 | entrezgene_transcript_name | EntrezGene transcript name ID |
| 60 | hpa | Human Protein Atlas Antibody ID |
| 61 | ottg | VEGA gene ID(s) (OTTG) |
| 62 | ottt | VEGA transcript ID(s) (OTTT) |
| 63 | ottp | VEGA protein ID(s) (OTTP) |
| 64 | hgnc_id | HGNC ID(s) |
| 65 | hgnc_symbol | HGNC symbol |
| 66 | hgnc_transcript_name | HGNC transcript name |
| 67 | merops | MEROPS ID |
| 68 | mim_morbid_accession | MIM Morbid Accession |
| 69 | mim_morbid_description | MIM Morbid Description |
| 70 | mim_gene_accession | MIM Gene Accession |
| 71 | mim_gene_description | MIM Gene Description |
| 72 | mirbase_accession | miRBase Accession(s) |
| 73 | mirbase_id | miRBase ID(s) |
| 74 | mirbase_transcript_name | miRBase transcript name |
| 75 | pdb | PDB ID |
| 76 | protein_id | Protein (Genbank) ID [e.g. AAA02487] |
| 77 | pubmed | PubMed ID [e.g. 7716543] |
| 78 | reactome | Reactome ID |
| 79 | reactome_gene | Reactome gene ID [e.g. REACT_1006] |
| 80 | reactome_transcript | Reactome transcript ID [e.g. REACT_11045] |
| 81 | refseq_mrna | RefSeq mRNA [e.g. NM_001195597] |
| 82 | refseq_mrna_predicted | RefSeq mRNA predicted [e.g. XM_001125684] |
| 83 | refseq_ncrna | RefSeq ncRNA [e.g. NR_002834] |
| 84 | refseq_ncrna_predicted | RefSeq ncRNA predicted [e.g. XR_108264] |
| 85 | refseq_peptide | RefSeq Protein ID [e.g. NP_001005353] |
| 86 | refseq_peptide_predicted | RefSeq Predicted Protein ID [e.g. XP_001720922] |
| 87 | rfam | Rfam ID |
| 88 | rfam_transcript_name | Rfam transcript name |
| 89 | rnacentral | RNACentral ID |

| | | |
|-----|--------------------------------------|--|
| 90 | ucsc | UCSC ID |
| 91 | unigene | Unigene ID |
| 92 | uniparc | UniParc |
| 93 | uniprot_sptrembl | UniProt/TrEMBL Accession |
| 94 | uniprot_swissprot | UniProt/SwissProt Accession |
| 95 | uniprot_genename | UniProt Gene Name |
| 96 | uniprot_genename_transcript_name | Uniprot Transcript Name |
| 97 | wikigene_name | WikiGene Name |
| 98 | wikigene_id | WikiGene ID |
| 99 | wikigene_description | WikiGene Description |
| 100 | efg_agilent_sureprint_g3_ge_8x60k | Agilent SurePrint G3 GE 8x60k probe |
| 101 | efg_agilent_sureprint_g3_ge_8x60k_v2 | Agilent SurePrint G3 GE 8x60k v2 probe |
| 102 | efg_agilent_wholegenome_4x44k_v1 | Agilent WholeGenome 4x44k v1 probe |
| 103 | efg_agilent_wholegenome_4x44k_v2 | Agilent WholeGenome 4x44k v2 probe |
| 104 | affy_hc_g110 | Affy HC G110 probeset |
| 105 | affy_hg_focus | Affy HG FOCUS probeset |
| 106 | affy_hg_u133_plus_2 | Affy HG U133-PLUS-2 probeset |
| 107 | affy_hg_u133a_2 | Affy HG U133A_2 probeset |
| 108 | affy_hg_u133a | Affy HG U133A probeset |
| 109 | affy_hg_u133b | Affy HG U133B probeset |
| 110 | affy_hg_u95av2 | Affy HG U95AV2 probeset |
| 111 | affy_hg_u95b | Affy HG U95B probeset |
| 112 | affy_hg_u95c | Affy HG U95C probeset |
| 113 | affy_hg_u95d | Affy HG U95D probeset |
| 114 | affy_hg_u95e | Affy HG U95E probeset |
| 115 | affy_hg_u95a | Affy HG U95A probeset |
| 116 | affy_hugenefl | Affy HuGene FL probeset |
| 117 | affy_hta_2_0 | Affy HTA-2_0 probeset |
| 118 | affy_huex_1_0_st_v2 | Affy HuEx 1_0 st v2 probeset |
| 119 | affy_hugene_1_0_st_v1 | Affy HuGene 1_0 st v1 probeset |
| 120 | affy_hugene_2_0_st_v1 | Affy HuGene 2_0 st v1 probeset |
| 121 | affy_primeview | Affy primeview |
| 122 | affy_u133_x3p | Affy U133 X3P probeset |
| 123 | agilent_cgh_44b | Agilent CGH 44b probe |
| 124 | codelink | Codelink probe |
| 125 | illumina_humanwg_6_v1 | Illumina HumanWG 6 v1 probe |
| 126 | illumina_humanwg_6_v2 | Illumina HumanWG 6 v2 probe |
| 127 | illumina_humanwg_6_v3 | Illumina HumanWG 6 v3 probe |
| 128 | illumina_humanht_12_v3 | Illumina Human HT 12 V3 probe |
| 129 | illumina_humanht_12_v4 | Illumina Human HT 12 V4 probe |
| 130 | illumina_humanref_8_v3 | Illumina Human Ref 8 V3 probe |
| 131 | phalanx_onearray | Phalanx OneArray probe |
| 132 | family | Ensembl Protein Family ID(s) |
| 133 | family_description | Ensembl Family Description |
| 134 | pirsf | PIRSF ID |

| | | |
|-----|----------------------------|------------------------------------|
| 135 | pirsf_start | PIRSF start |
| 136 | pirsf_end | PIRSF end |
| 137 | superfamily | SUPERFAMILY ID |
| 138 | superfamily_start | SUPERFAMILY start |
| 139 | superfamily_end | SUPERFAMILY end |
| 140 | smart | SMART ID |
| 141 | smart_start | SMART start |
| 142 | smart_end | SMART end |
| 143 | hamap | HAMAP Accession ID |
| 144 | hamap_start | HAMAP start |
| 145 | hamap_end | HAMAP end |
| 146 | profile | Pfscan ID |
| 147 | profile_start | Pfscan start |
| 148 | profile_end | Pfscan end |
| 149 | prosite | ScanProsite ID |
| 150 | prosite_start | ScanProsite start |
| 151 | prosite_end | ScanProsite end |
| 152 | prints | PRINTS ID |
| 153 | prints_start | PRINTS start |
| 154 | prints_end | PRINTS end |
| 155 | pfam | Pfam ID |
| 156 | pfam_start | Pfam start |
| 157 | pfam_end | Pfam end |
| 158 | tigrfam | TIGRFAM ID |
| 159 | tigrfam_start | TIGRFAM start |
| 160 | tigrfam_end | TIGRFAM end |
| 161 | gene3d | Gene3D ID |
| 162 | gene3d_start | Gene3D start |
| 163 | gene3d_end | Gene3D end |
| 164 | hmmpanther | HMMPanther ID |
| 165 | hmmpanther_start | HMMPanther start |
| 166 | hmmpanther_end | HMMPanther end |
| 167 | interpro | Interpro ID |
| 168 | interpro_short_description | Interpro Short Description |
| 169 | interpro_description | Interpro Description |
| 170 | interpro_start | Interpro start |
| 171 | interpro_end | Interpro end |
| 172 | low_complexity | low complexity (SEG) |
| 173 | low_complexity_start | low complexity (SEG) start |
| 174 | low_complexity_end | low complexity (SEG) end |
| 175 | transmembrane_domain | Transmembrane domain (tmhmm) |
| 176 | transmembrane_domain_start | Transmembrane domain (tmhmm) start |
| 177 | transmembrane_domain_end | Transmembrane domain (tmhmm) end |
| 178 | signal_domain | signal peptide |
| 179 | signal_domain_start | signal peptide start |

```

180             signal_domain_end                signal peptide end
181             ncoils                          coiled coil (ncoils)
182             ncoils_start                    coiled coil (ncoils) start
183             ncoils_end                      coiled coil (ncoils) end

```

We now get a short list of attributes related to the region where the genes are located.

8 Local BioMart databases

The biomaRt package can be used with a local install of a public BioMart database or a locally developed BioMart database and web service. In order for biomaRt to recognize the database as a BioMart, make sure that the local database you create has a name conform with

```
database_mart_version
```

where database is the name of the database and version is a version number. No more underscores than the ones showed should be present in this name. A possible name is for example

```
ensemblLocal_mart_46
```

8.1 Minimum requirements for local database installation

More information on installing a local copy of a BioMart database or develop your own BioMart database and webservice can be found on <http://www.biomart.org> Once the local database is installed you can use biomaRt on this database by:

```
listMarts(host="www.myLocalHost.org", path="/myPathToWebservice/martservice")
mart=useMart("nameOfMyMart",dataset="nameOfMyDataset",host="www.myLocalHost.org", path="/myPathToWebservice/martser
```

For more information on how to install a public BioMart database see: <http://www.biomart.org/install.html> and follow link databases.

9 Using select

In order to provide a more consistent interface to all annotations in Bioconductor the `select`, `columns`, `keytypes` and `keys` have been implemented to wrap some of the existing functionality above. These methods can be called in the same manner that they are used in other parts of the project except

that instead of taking a `AnnotationDb` derived class they take instead a `Mart` derived class as their 1st argument. Otherwise usage should be essentially the same. You still use `columns` to discover things that can be extracted from a `Mart`, and `keytypes` to discover which things can be used as keys with `select`.

```
> mart<-useMart(dataset="hsapiens_gene_ensembl",biomart='ensembl')
> head(keytypes(mart), n=3)

[1] "affy_hc_g110"          "affy_hg_focus"        "affy_hg_u133_plus_2"
> head(columns(mart), n=3)

[1] "3_utr_end"    "3_utr_end"    "3_utr_start"
```

And you still can use `keys` to extract potential keys, for a particular key type.

```
> k = keys(mart, keytype="chromosome_name")
> head(k, n=3)

[1] "1" "2" "3"
```

When using `keys`, you can even take advantage of the extra arguments that are available for others keys methods.

```
> k = keys(mart, keytype="chromosome_name", pattern="LRG")
> head(k, n=3)

character(0)
```

Unfortunately the `keys` method will not work with all key types because they are not all supported.

But you can still use `select` here to extract columns of data that match a particular set of keys (this is basically a wrapper for `getBM`).

```
> affy=c("202763_at","209310_s_at","207500_at")
> select(mart, keys=affy, columns=c('affy_hg_u133_plus_2','entrezgene'),
+   keytype='affy_hg_u133_plus_2')

  affy_hg_u133_plus_2  entrezgene
1      209310_s_at      837
2      207500_at      838
3      202763_at      836
```

So why would we want to do this when we already have functions like `getBM`? For two reasons: 1) for people who are familiar with `select` and its helper methods, they can now proceed to use `biomaRt` making the same kinds of calls that are already familiar to them and 2) because the `select` method is implemented in many places elsewhere, the fact that these methods are shared allows for more convenient programmatic access of all these resources. An example of a package that takes advantage of this is the *OrganismDbi* package. Where several packages can be accessed as if they were one resource.

10 Session Info

```
> sessionInfo()
```

```
R version 3.2.2 (2015-08-14)
Platform: x86_64-pc-linux-gnu (64-bit)
Running under: Ubuntu 14.04.3 LTS
```

```
locale:
```

```
[1] LC_CTYPE=en_US.UTF-8      LC_NUMERIC=C               LC_TIME=en_US.UTF-8
[5] LC_MONETARY=en_US.UTF-8   LC_MESSAGES=en_US.UTF-8   LC_PAPER=en_US.UTF-8
[9] LC_ADDRESS=C              LC_TELEPHONE=C            LC_MEASUREMENT=en_US.UTF-8
```

```
attached base packages:
```

```
[1] stats      graphics  grDevices  utils      datasets  methods    base
```

```
other attached packages:
```

```
[1] biomaRt_2.26.1
```

```
loaded via a namespace (and not attached):
```

```
[1] IRanges_2.4.4      parallel_3.2.2      DBI_0.3.1           tools_3.2.2
[6] Biobase_2.30.0     AnnotationDbi_1.32.1 RSQLite_1.0.0       S4Vectors_0.8.3
[11] stats4_3.2.2       bitops_1.0-6        XML_3.98-1.3
```

```
> warnings()
```

```
NULL
```