Package ‘InteractionSet’

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**Description**

Computing a minimum bounding box for groups of pairwise interactions.

**Usage**

```r
## S4 method for signature 'GInteractions'
boundingBox(x, f)
```

# Note, the same call is used for the InteractionSet method.

**Arguments**

- `x`: A GInteractions or InteractionSet object.
- `f`: A factor or vector of length equal to that of `x`, indicating the group to which each pairwise interaction belongs.

**Details**

For any group of pairwise interactions, the minimum bounding box is the smallest rectangle in the interaction space that contains all interactions in the group. Each side of the box has coordinates spanning the most extreme anchor regions on the corresponding chromosome. This is often useful for summarizing clusters of interactions.

Grouping of interactions is specified using `f`, where interactions in `x` with the same level of `f` are considered to be in the same group. If `f` is not specified, all interactions in `x` are assumed to be in a single group (named as “1”). An error will be raised if a group spans multiple chromosomes for either the first or second anchor regions.

The function returns a GInteractions object containing the coordinates of the bounding boxes for all groups. Each interaction represents a bounding box for a group, where the anchor regions represent the sides of the box. Entries are named according to the levels of `f`, in order to specify which bounding box corresponds to which group.

It is recommended to run `swapAnchors` prior to computing the bounding box for intra-chromosomal groups. If all `anchor1 >= anchor2` or all `anchor1 <= anchor2`, all interactions will lie on one side of the diagonal of the intra-chromosomal interaction space. This results in the smallest possible minimum bounding box, which will only increase in size if interactions are placed on the other side of the diagonal. Alternatively, users can specify a StrictGInteractions object as an input into `x`, in which `anchor1 <= anchor2` is enforced automatically.
Value

A GInteractions object containing the coordinates of each bounding box.

Author(s)

Aaron Lun

See Also

GInteractions-class, InteractionSet-class

Examples

```r
example(GInteractions, echo=FALSE)
# Making up a sensible grouping.
gi <- sort(gi)
all.chrs <- as.character(seqnames(regions(gi)))
f <- paste0(all.chrs[anchors(gi, type="first", id=TRUE)], ".",
             all.chrs[anchors(gi, type="second", id=TRUE)])

boundingBox(gi, f)
boundingBox(swapAnchors(gi), f)
# Fails for multiple chromosomes
try(out <- boundingBox(gi))
in.A <- f=="chrA.chrA"
out <- boundingBox(gi[in.A])
```

Description

Methods to get and set fields in a ContactMatrix object.

Usage

```r
## S4 method for signature 'ContactMatrix'
anchors(x, type="both", id=FALSE)
## S4 replacement method for signature 'ContactMatrix'
anchors(x) <- value

## S4 method for signature 'ContactMatrix'
regions(x)
## S4 replacement method for signature 'ContactMatrix'
regions(x) <- value

## S4 method for signature 'ContactMatrix'
replaceRegions(x) <- value
## S4 replacement method for signature 'ContactMatrix'
appendRegions(x) <- value
```
ContactMatrix accessors

## S4 method for signature 'ContactMatrix'
reduceRegions(x)

## S4 method for signature 'ContactMatrix'
show(x)

## S4 method for signature 'ContactMatrix'
as.matrix(x)

## S4 replacement method for signature 'ContactMatrix'
as.matrix(x) <- value

## S4 method for signature 'ContactMatrix'
t(x)

dim(x)

dimnames(x)

## S4 replacement method for signature 'ContactMatrix'
dimnames(x) <- value

## S4 method for signature 'ContactMatrix'
length(x)

### Arguments

- **x**: A ContactMatrix object.
- **type**: a string specifying which anchors are to be extracted or replaced.
- **id**: a scalar logical indicating whether indices or GRanges should be returned.
- **value**: For anchors<- and type="row" or "column", an integer vector of length equal to the number of rows or columns in x, respectively. For type="both", a list of two such vectors must be supplied (first and second for rows and columns, respectively).
  
  For anchors<-, a list of two integer vectors when type="both". The first and second vectors must have length equal to the number of rows and columns of x, respectively. For type="row" or "column", only one vector needs to be supplied (rows or columns, respectively).
  
  For regions<-, a GRanges object of length equal to that of regions(x). For newRegions<-, a GRanges object that is a superset of all entries in regions(x) involved in interactions. For appendRegions<-, a GRanges of any length containing additional regions.
  
  For as.matrix<-, a Matrix object of the same dimensions as that in the matrix slot.
  
  For dimnames<-, a list of two character vectors corresponding to the row and column names, respectively. These can also be passed separately via rownames<- and colnames<-

### Details

The return value of anchors varies depending on type and id. If id=FALSE, a GRanges object will be returned containing the regions specified by the anchor1 or anchor2 slots in x, for type="row" or "column", respectively. If type="both", a GRangesList will be returned with two entries - row and column, containing regions specified by anchor1 and anchor2 respectively. However, if id=TRUE, the integer vectors in the anchor1 or anchor2 slots of x are returned directly (a list of length two is returned for type="both", containing both of the vectors).
Replacement in anchors<- can only be performed using anchor indices. If type="both", a list of two integer vectors is required in value, specifying the indices of the row- and column-wise interacting regions in x. If type="row" or "column", an integer vector is required to replace the existing row- or column-wise indices in the anchor1 or anchor2 slot, respectively.

For regions, a GRanges is returned equal to the regions slot in x. For regions<-, the GRanges in value is used to replace the regions slot. Resorting of the replacement GRanges is performed automatically, along with re-indexing of the anchors. In addition, the input GRanges must be of the same length as the existing object in the regions slot. The newRegions replacement method can take variable length GRanges, but requires that the replacement contain (at least) all ranges contained in anchors(x). The appendRegions replacement method appends extra intervals to the existing regions slot of x. The reduceRegions method removes unused entries in the regions slot, to save memory – see reduceRegions,GInteractions-method for more details.

The show method will print out various details of the object, such as the dimensions of the matrix slot and the length of the regions slot. The as.matrix method will return the value of the matrix slot, containing the Matrix of interaction data. Replacement with a Matrix of the same dimensions can be performed using the as.matrix<- function. The t method will transpose the matrix, i.e., switch the rows and columns (and switch the vectors in the anchor1 and anchor2 slots).

The dim method will return a vector of length 2, containing the dimensions of the matrix slot. The dimnames method will return a list of length 2, containing the row and column names of matrix (these can be modified with the dimnames<- method). The length method will return an integer scalar corresponding to the total number of entries in the matrix slot.

As the ContactMatrix class inherits from the Annotated class, additional metadata can be stored in the metadata slot. This can be accessed or modified with metadata,Annotated-method.

Value

For the getters, values in various slots of x are returned, while for the setters, the slots of x are modified accordingly – see Details.

Author(s)

Aaron Lun

See Also

ContactMatrix-class

Examples

eexample(ContactMatrix, echo=FALSE) # Generate a nice object.
show(x)

# Various matrix methods:
as.matrix(x)
t(x)
nrow(x)
ncol(x)
length(x)

# Accessing anchor ranges or indices:
anchors(x)
anchors(x, type="row")
ContactMatrix class

### Description

The ContactMatrix class contains a matrix where rows and columns represent genomic loci. Each entry of the matrix contains information about the interaction between the loci represented by the corresponding row/column, e.g., contact frequencies. Coordinates of the loci are also contained within this class.

### Usage

```r
# S4 method for signature 'ANY, numeric, numeric, GRanges'
ContactMatrix(matrix, anchor1, anchor2, regions, metadata=list())

# S4 method for signature 'ANY, GRanges, GRanges, GenomicRangesORmissing'
ContactMatrix(matrix, anchor1, anchor2, regions, metadata=list())

# S4 method for signature 'missing, missing, missing, GenomicRangesORmissing'
ContactMatrix(matrix, anchor1, anchor2, regions, metadata=list())
```
ContactMatrix class

Arguments

matrix
Any object containing interaction data that can be coerced into a Matrix object.

anchor1, anchor2
Either a pair of numeric vectors containing indices to regions or a pair of GRanges objects. In both cases, anchor1 and anchor2 should have lengths equal to the number of rows and columns in matrix, respectively.

regions
A GRanges object containing the coordinates of the interacting regions. This argument is optional for InteractionSet, ANY, GRanges, GRanges-method.

metadata
A list containing experiment-wide metadata - see ?Annotated for more details.

Details

The ContactMatrix class inherits from the Annotated class, with several additional slots:

matrix: A Matrix object.
anchor1: An integer vector specifying the index of the first interacting region.
anchor2: An integer vector specifying the index of the second interacting region.
regions: A sorted GRanges object containing the coordinates of all interacting regions.

Each entry of anchor1 corresponds to a row in matrix, while each entry of anchor2 corresponds to a column. Each entry of matrix represents an interaction between the corresponding entries in anchor1 and anchor2, which point to the relevant coordinates in regions for each locus.

ContactMatrix objects can be constructed by specifying numeric vectors as anchor1 and anchor2 in the ContactMatrix function. These vectors will define the regions corresponding to the rows and columns of the matrix. Specifically, each value of the vector acts as an index to specify the relevant coordinates from regions. This means that the range of entries must lie within [1, length(regions)].

Alternatively, ContactMatrix objects can be constructed by directly supplying the GRanges of the interacting loci in ContactMatrix. If regions is not specified, this will be constructed automatically from the two sets of supplied GRanges. If regions is supplied, exact matching will be performed to identify the indices in regions corresponding to the regions in the supplied GRanges. Missing values are not tolerated and will cause an error to be raised.

Both methods will return an ContactMatrix object containing all of the specified information. Sorting of regions is also performed automatically, with re-indexing of all anchor indices to preserve the correct pairings between regions.

Value

For the constructors, a ContactMatrix object is returned.

Choosing between Matrix classes

The ContactMatrix class provides support for Matrix objects from the Matrix package. This provides several classes for storing sparse and symmetric matrices in a memory-efficient manner, e.g., via the sparseMatrix class. These classes may be preferable for handling very large matrices spanning substantial areas in the interaction space.

There are, however, some caveats to the use of the various Matrix subclasses. All numeric values are coerced to double-precision types, which may take up more memory than a direct integer representation. Character data is not supported, though this is unlikely to be relevant in most cases. Another issue is how missing values should be interpreted in the sparseMatrix – see ?inflate for more details.
In general, the choice of class depends on the type of data and the intended application. A dense Matrix is often sufficient for most use cases where small or dense regions of the interaction space are being examined. Switching to sparseMatrix may be desirable in emptier parts of the space, where there can be substantial memory savings.

Author(s)

Aaron Lun

See Also

ContactMatrix-access, ContactMatrix-subset, ContactMatrix-sort, Annotated-class, Matrix-class

Examples

```r
set.seed(1000)
N <- 30
all.starts <- round(runif(N, 1, 100))
all.ends <- all.starts + round(runif(N, 5, 20))
all.regions <- GRanges(rep(c("chrA", "chrB"), c(N-10, 10)),
                      IRanges(all.starts, all.ends))
Nr <- 10
Nc <- 20
all.anchor1 <- sample(N, Nr)
all.anchor2 <- sample(N, Nc)
counts <- matrix(rpois(Nr*Nc, lambda=10), Nr, Nc)
x <- ContactMatrix(counts, all.anchor1, all.anchor2, all.regions)
# Equivalent construction:
x <- ContactMatrix(counts, all.regions[all.anchor1],
                   all.regions[all.anchor2])
x <- ContactMatrix(counts, all.regions[all.anchor1],
                   all.regions[all.anchor2], all.regions)
# Also works directly with Matrix objects.
counts2 <- Matrix::Matrix(counts)
x2 <- ContactMatrix(counts2, all.anchor1, all.anchor2, all.regions)
counts2 <- as(counts2, "dgCMatrix")
x2 <- ContactMatrix(counts2, all.anchor1, all.anchor2, all.regions)
```

ContactMatrix distances

*Compute linear distances from ContactMatrix objects*

Description

Methods to compute linear distances between pairs of interacting regions in a ContactMatrix object.
ContactMatrix distances

Usage

```r
## S4 method for signature 'ContactMatrix'
pairdist(x, type="mid")
```

```r
## S4 method for signature 'ContactMatrix'
intrachr(x)
```

Arguments

- **x**: A ContactMatrix object.
- **type**: A character string specifying the type of distance to compute. See `pairdist,InteractionSet-method` for possible values.

Details

`pairdist,ContactMatrix-method` will return a matrix of integer (or, if `type="intra"`, logical) values. Each entry of this matrix specifies the distance between the interacting loci that are represented by the corresponding row and column. If `type="intra"`, each entry indicates whether the corresponding interaction is intra-chromosomal. Running `intrachr(x)` is equivalent to `pairdist(x, type="intra")` for any ContactMatrix object `x`. See `pairdist,InteractionSet-method` for more details on the type of distances that can be computed.

Value

An integer or logical matrix of the same dimensions as `x`, containing the specified distances.

Author(s)

Aaron Lun

See Also

`ContactMatrix-class,pairdist,InteractionSet-method`

Examples

```r
example(ContactMatrix, echo=FALSE)
pairdist(x)
pairdist(x, type="gap")
pairdist(x, type="span")
pairdist(x, type="diag")
intrachr(x)
```
ContactMatrix overlaps

Find overlaps between GRanges and a ContactMatrix

Description

Find overlaps between a set of linear intervals in a GRanges object, and the set of regions representing the rows or columns of a ContactMatrix.

Usage

## S4 method for signature 'ContactMatrix,GRanges'
overlapsAny(query, subject, maxgap=0L, minoverlap=1L,
type=c("any", "start", "end", "within", "equal"),
ignore.strand=TRUE)

## S4 method for signature 'ContactMatrix,GRangesList'
overlapsAny(query, subject, maxgap=0L, minoverlap=1L,
type=c("any", "start", "end", "within", "equal"),
ignore.strand=TRUE, use.region="both")

# The same call is used for the methods taking GRangesList, GInteractions and # InteractionSet objects as 'subject'. For brevity, these will not be listed.

Arguments

query A ContactMatrix object.
subject A GRanges, GRangesList, GInteractions or InteractionSet object.
maxgap, minoverlap, type
See ?findOverlaps in the GenomicRanges package.
ignore.strand See ?findOverlaps in the GenomicRanges package.
use.region A string specifying how the interacting regions are to be matched to row/column regions.

Details

When subject is a GRanges, overlaps are identified between the row regions of query and the regions in subject. This is repeated for the column regions of query. A list of two logical vectors is returned, specifying the rows and columns in query that are overlapped by any region in subject. These vectors can be directly used to subset query. Alternatively, they can be used in outer to generate a logical matrix for masking – see Examples.

For all other classes of subject, two-dimensional overlaps are identified. A logical matrix is returned indicating which entries in the ContactMatrix have overlaps with the specified interactions. For any given entry, an overlap is only considered if the regions for the corresponding row and column both overlap anchor regions in the subject. See ?"findOverlaps,GInteractions,GInteractions-method" for more details.

If use.region="both", overlaps between any row/column region and the first/second interacting region of subject are considered. If use.region="same", only overlaps between row regions and the first interacting regions, or between column regions and the second interacting regions are considered. If use.region="reverse", only overlaps between row regions and the second interacting regions, or between row regions and the first interacting regions are considered.
Value

For \texttt{overlapsAny,ContactMatrix,GRanges-method}, a named list of two logical vectors is returned specifying the rows of columns of \texttt{query} overlapped by \texttt{subject}.

For the other \texttt{overlapsAny} methods, a logical matrix is returned indicating which entries in \texttt{query} are overlapped by \texttt{subject}.

Author(s)

Aaron Lun

See Also

\texttt{ContactMatrix-class,findOverlaps}

Examples

```r
example(ContactMatrix, echo=FALSE)

of.interest <- resize(sample(regions(x), 2), width=1, fix="center")
olap <- overlapsAny(x, of.interest)
olap
x[olap$row,]
x[,olap$column]
x[olap$row,olap$column]

keep <- outer(olap$row, olap$column, "|") # OR mask
temp <- as.matrix(x)
temp[!keep] <- NA

keep <- outer(olap$row, olap$column, ";") # AND mask
temp <- as.matrix(x)
temp[!keep] <- NA

# Two dimensional overlaps.
pairing <- GRangesList(first=regions(x), second=regions(x))
olap <- overlapsAny(x, pairing)
olap
olap <- overlapsAny(sort(x), pairing) # A bit prettier
olap
```
ContactMatrix sorting

Usage

```r
## S4 method for signature 'ContactMatrix'
order(..., na.last=TRUE, decreasing=FALSE)
## S4 method for signature 'ContactMatrix'
sort(x, decreasing=FALSE, ...)
## S4 method for signature 'ContactMatrix'
duplicated(x, incomparables=FALSE, fromLast=FALSE, ...)
## S4 method for signature 'ContactMatrix'
unique(x, incomparables=FALSE, fromLast=FALSE, ...)
```

Arguments

... For `sort`, `ContactMatrix-method`, one or more `ContactMatrix` objects with the same dimensions. Otherwise, ignored in all other methods.

x A `ContactMatrix` object.

na.last A logical scalar indicating whether NA values should be ordered last. This should not be relevant as anchor indices should be finite.

decreasing A logical scalar indicating whether sorting should be performed in decreasing order.

incomparables A logical scalar, ignored.

fromLast A logical scalar indicating whether the last entry of a repeated set in `x` should be considered as a non-duplicate.

Details

Sorting is performed based on the anchor indices of the `ContactMatrix` object. Rows are ordered for increasing values of the anchor1 slot, while columns are ordered for increasing values of the anchor2 slot. This equates to ordering by the coordinates directly, as the GRanges in the regions slot is always sorted. Based on this, `sort`, `ContactMatrix-method` will return a sorted copy of `x` with permuted rows/columns in increasing order. This can be set to decreasing order by specifying `decreasing=TRUE`.

`order`, `ContactMatrix-method` returns a list of 2 integer vectors. The first vector contains the permutation to rearrange the rows of `x` in increasing order, while the second vector does the same for the columns of `x`. If multiple objects are supplied to `order`, ordering will be computed using anchor indices from successive objects. In other words, ordering will be performed using anchor indices from the first object; any rows with the same anchor1 or columns with the same anchor2 will be split using the corresponding indices in the second object; and so on.

`duplicated`, `ContactMatrix-method` returns a list of two logical vectors. The first vector indicates whether rows are duplicated, based on identical values in the anchor1 slot. The second does the same for columns based on the anchor2 slot. For a set of duplicated rows or columns, the first occurrence of that row/column is marked as the non-duplicate if `fromLast=FALSE`, and the last entry otherwise.

`unique`, `ContactMatrix-method` returns a `ContactMatrix` object where all duplicate rows and columns have been removed from `x`. This is equivalent to subsetting based on the non-duplicate rows and columns defined in `duplicated`, `ContactMatrix-method`.

Note that sorting and duplicate identification only use the anchor indices. The values of the matrix slot are not used in distinguishing between rows or columns with the same index.
Value

For `sort` and `unique`, a `ContactMatrix` object is returned with sorted or unique rows/columns, respectively.

For `order`, a list of two integer vectors is returned containing row/column permutations.

For `duplicated`, a list of logical vectors is returned specifying which rows/columns are duplicated.

Author(s)

Aaron Lun

See Also

`ContactMatrix-class`

Examples

```r
example(ContactMatrix, echo=FALSE)

anchors(x)

x2 <- sort(x)

anchors(x2)

# Can also order them.
o <- order(x)

stopifnot(all.equal(x[o$row, o$column], x2))

# Checking duplication.
duplicated(x)

duplicated(rbind(x, x))

stopifnot(all.equal(x, unique(rbind(x, x))))
```

ContactMatrix subsetting

Methods to subset or combine `ContactMatrix` objects.

Usage

### Subsetting

```r
## S4 method for signature 'ContactMatrix,ANY,ANY'
x[i, j, ..., drop=TRUE]
## S4 replacement method for signature 'ContactMatrix,ANY,ANY,ContactMatrix'
x[i, j] <- value
## S4 method for signature 'ContactMatrix'
subset(x, i, j)
```
### Combining

#### S4 method for signature 'ContactMatrix'
```
cbind(..., deparse.level=1)
```
#### S4 method for signature 'ContactMatrix'
```
rbind(..., deparse.level=1)
```

**Arguments**

- **x**: A ContactMatrix object.
- **i, j**: A vector of subscripts, indicating the rows and columns to be subsetted for i and j, respectively.
- **...**: For `cbind`, `rbind` and `c`, ... contains ContactMatrix objects to be combined. Otherwise, this argument is ignored during subsetting.
- **drop**: A logical scalar, ignored by `[,ContactMatrix,ANY,ANY-method`.
- **value**: A ContactMatrix object with dimensions equal to the length of the two subscripts (or the corresponding dimensions of `x`, if either subscript is missing.
- **deparse.level**: An integer scalar; see `?base::cbind` for a description of this argument.

**Details**

Subsetting of ContactMatrix objects behaves like that for standard matrices. Rows and columns can be selected and rearranged, with concomitant changes to the anchor1 and anchor2 slots. All subsetting operations will return an ContactMatrix with the specified rows and columns. However, note that the value of regions will not be modified by subsetting.

cbind will combines objects with the same rows but different columns. Errors will occur if the row regions are not identical between objects (i.e., must have same values in the slots for regions and anchor1). Conversely, rbind will combines objects with the same columns but different rows. Again, errors will occur if the columns are not identical (i.e., must have same values in the slots for regions and anchor2).

If subsetted anchors are required, see "interaction-subset" for why subsetting should be done before calling the anchors method.

**Value**

A subsetted or combined ContactMatrix object.

**Author(s)**

Aaron Lun

**See Also**

ContactMatrix-class

**Examples**

```
example(ContactMatrix, echo=FALSE)
```

# Subsetting:
```
x[1:5,]
x[,10:15]
```
```r
x[1:5,10:15]
x2 <- x
x2[1:5,] <- x[6:10,]
as.matrix(x2[,1]) <- 20

# Combining
cbind(x, x)
rbind(x, x)
```

---

Convert classes

### Description
Inflate a GInteractions or InteractionSet into a ContactMatrix, or deflate a ContactMatrix to an InteractionSet.

### Usage

#### S4 method for signature 'GInteractions'
```r
inflate(x, rows, columns, fill, swap=TRUE, sparse=FALSE, ...)
```

#### S4 method for signature 'InteractionSet'
```r
inflate(x, rows, columns, assay=1L, sample=1L, fill=NULL, swap=TRUE, sparse=FALSE, ...)
```

#### S4 method for signature 'ContactMatrix'
```r
deflate(x, collapse=TRUE, extract, use.zero, use.na, ...)
```

### Arguments

- **x**: A GInteractions or InteractionSet object for inflate, or a ContactMatrix object for deflate.
- **rows, columns**: An integer, logical or character vector, a GRanges object or NULL, indicating the regions of interest to be used as the rows or columns of the ContactMatrix.
- **assay**: A numeric scalar indicating the assay of the InteractionSet object, from which values are extracted to fill the ContactMatrix.
- **sample**: A numeric scalar indicating the sample (i.e., column) of the assay to extract values to fill the ContactMatrix.
- **fill**: A numeric vector of length equal to nrow(x), containing values with which to fill the ContactMatrix. If specified, this overrides extraction of assay values for inflate, InteractionSet-method.
- **swap**: A logical scalar indicating whether filling should also be performed after swapping anchor indices.
- **sparse**: A logical scalar indicating whether the inflated matrix should use a sparseMatrix representation.
- **collapse**: A logical scalar indicating whether duplicated interactions should be removed from x prior to deflation.
- **extract**: A logical vector or matrix indicating which entries of x to convert into pairwise interactions.
Convert classes

use.zero, use.na
A logical scalar indicating whether to convert zero or NA entries into pairwise interactions.

For inflate, additional arguments to pass to overlapsAny when rows or columns is a GRanges.
For deflate, additional arguments to pass to the InteractionSet constructor.

Value
For inflate, a ContactMatrix is returned.
For deflate, an InteractionSet object is returned.

Inflating to a ContactMatrix

The inflate method will return a ContactMatrix where the rows and columns correspond to specified regions of interest in rows and columns. Regions can be specified by supplying an object of various types:

- If it is an integer vector, it is assumed to refer to intervals in the regions slot of the input object x. Values of the vector need not be sorted or unique, but must lie within [1, regions(x)].
- If it is a logical vector, it will subset to retain intervals in regions(x) that are TRUE.
- If it is a character vector, it is assumed to contain the names of the reference sequences of interest (i.e., chromosome names).
- If it is a GRanges object, overlapsAny will be called to identify the overlapping intervals of regions(x).
- If it is NULL, all regions in regions(x) will be used to construct that dimension of the ContactMatrix.

For the GIInteractions method, values in the matrix are filled based on user-supplied values in fill. Each entry of fill corresponds to an interaction in x is used to set the matrix value at the matching row/column. Some entries of the matrix will correspond to pairwise interactions that are not present in x - these are filled with NA values.

By default, filling is repeated after swapping the anchor indices. This means that the value of the matrix at (1, 2) will be the same as that at (2, 1), i.e., the matrix is symmetric around the diagonal of the interaction space. However, if swap=FALSE, filling is performed so that the first and second anchor indices correspond strictly to rows and columns, respectively. This may be preferable if the order of the anchors contains some relevant information. In all cases, if duplicated interactions are present in x (and redundant permutations, when swap=TRUE), one will be arbitrarily chosen to fill the matrix.

For the InteractionSet inflate method, entries in the matrix are filled in based on the values in the first sample of the first assay by default. For more complex x, values from different assays and samples can be extracted using the assay and sample arguments. Note that if fill is specified, it will override any extraction of values from the assays.

If sparse=TRUE, inflate will return a ContactMatrix containing a sparseMatrix in the matrix slot. Here, entries without a corresponding interaction in x are set to zero, not NA.

Deflating from a ContactMatrix

The deflate method will return an InteractionSet where each relevant entry in the ContactMatrix is converted into a pairwise interaction. Relevant entries are defined as those that are non-zero, if use.zero is FALSE; and non-NA, if use.na is FALSE. If x contains a sparseMatrix representation,
the former is set to FALSE while the latter is set to TRUE, if either are not specified. For all other matrices, use.zero=TRUE and use.na=FALSE by default.

If extract is specified, this overrides all values of use.zero and use.na. A typical application would be to deflate a number of ContactMatrix objects with the same extract matrix. This ensures that the resulting InteractionSet objects can be easily combined with cbind, as the interactions are guaranteed to be the same. Otherwise, different interactions may be extracted depending on the presence of zero or NA values.

The values of all matrix entries are stored as a one-sample assay, with each value corresponding to its pairwise interaction after conversion. Duplicate interactions are removed by default, along with redundant permutations of the anchor indices. These can be included in the returned object by setting collapse=FALSE. Additional arguments can be used to specify the colData and metadata, which are stored in the ContactMatrix itself.

### Interpreting zeroes in a sparse matrix

Storing data as a sparseMatrix may be helpful as it is more memory-efficient for sparse areas of the interaction space. However, users should keep in mind that the zero values in the sparseMatrix may not represent zeroes in fill. The majority of these values are likely to be zero just because there was no corresponding interaction in x to set it to a non-zero value.

Whether or not this is a problem depends on the application. For example, if fill represents count data and only interactions with non-zero counts are stored in x, then setting all other entries to zero is sensible. However, in other cases, it is not appropriate to fill entries corresponding to missing interactions with zero. If fill represents, e.g., log-fold changes, then setting missing entries to a value of zero will be misleading. One could simply ignore zeroes altogether, though this will also discard entries that are genuinely zero.

These problems are largely avoided with the default denseMatrix classes, where missing entries are simply set to NA.

### Author(s)

Aaron Lun

### See Also

*InteractionSet-class, GInteractions-class, ContactMatrix-class, sparseMatrix-class*

### Examples

```r
example(InteractionSet, echo=FALSE)
inflate(iset, 1:10, 1:10)
inflate(iset, 1:10, 1:10, sparse=TRUE)
inflate(iset, 1:10, 1:5+10)
inflate(iset, "chrA", 1:5+10)
inflate(iset, "chrA", "chrB")
inflate(iset, "chrA", GRanges("chrB", IRanges(1, 10)))

y <- inflate(iset, 1:10, 1:10)
iset2 <- deflate(y)
iset2
 assays(iset2)

y <- inflate(iset, 1:10, 1:10, swap=FALSE)
```
iset2 <- deflate(y)
iset2
assay(iset2)

# Testing with different fillings:
y <- inflate(iset, 1:10, 1:10, sample=2)
iset2 <- deflate(y)
assay(iset2)

y <- inflate(iset, 1:10, 1:10, fill=rowSums(assay(iset)))
iset2 <- deflate(y)
assay(iset2)

y2 <- inflate(interactions(iset), 1:10, 1:10, rowSums(assay(iset)))
identical(y, y2) # should be TRUE

# Effect of 'collapse'
y <- inflate(iset, c(8, 1:10), 1:10)
deflate(y)
deflate(y, collapse=FALSE)

---

### Description

The `GInteractions` class stores pairwise genomic interactions, and is intended for use in data analysis from Hi-C or ChIA-PET experiments. Each row of the `GInteractions` corresponds to a pairwise interaction between two loci, with indexing to improve computational efficiency.

### Usage

```r
## S4 method for signature 'numeric,numeric,GRanges'
GInteractions(anchor1, anchor2, regions, metadata=list(), mode="normal", ...)

## S4 method for signature 'GRanges,GRanges,GenomicRangesORmissing'
GInteractions(anchor1, anchor2, regions, metadata=list(), mode="normal", ...)

## S4 method for signature 'missing,missing,GenomicRangesORmissing'
GInteractions(anchor1, anchor2, regions, metadata=list(), mode="normal", ...)
```

### Arguments

- `anchor1, anchor2`
  - Either a pair of numeric vectors containing indices to `regions`, or a pair of `GRanges` objects specifying the interacting loci. Lengths of both `anchor1` and `anchor2` must be equal.

- `regions`
  - A `GRanges` object containing the coordinates of the interacting regions. This is only mandatory if `anchor1` and `anchor2` are numeric vectors.

- `metadata`
  - An optional list of arbitrary content describing the overall experiment.

- `mode`
  - A string indicating what type of `GInteractions` object should be constructed.

- `...`
  - Optional metadata columns.
GInteractions class

Value

For the constructors, a GInteractions (or StrictGInteractions, or ReverseStrictGInteractions) object is returned.

Description of the class

The GInteractions class inherits from the Vector class and has access to all of its data members and methods (e.g., metadata and elementMetadata - see Vector-class for more details). It also contains several additional slots:

- anchor1: An integer vector specifying the index of the first interacting region.
- anchor2: An integer vector specifying the index of the second interacting region.
- regions: A sorted GRanges object containing the coordinates of all interacting regions.

Each interaction is defined by the corresponding entries in the anchor1 and anchor2 slots, which point to the relevant coordinates in regions for each locus.

The StrictGInteractions class inherits from the GInteractions class and has the same slots. The only difference is that, for each interaction, anchor1 must be less than or equal to anchor2. This means that the first interacting region has a start position that is "lower" than the second interacting region. This condition is useful for comparing interactions within and between objects, as it ensures that redundant permutations of the same interaction are not being overlooked. However, it is not used by default as there may conceivably be instances where the order of interactions is informative. The ReverseStrictGInteractions class has the opposite behaviour, where anchor1 must be greater than or equal to anchor2.

Class construction

GInteractions objects can be constructed by specifying integer vectors to define the pairwise interactions in the GInteractions call. For entry `x`, the corresponding interaction is defined between `regions[anchor1[x]]` and `regions[anchor2[x]]`. Obviously, coordinates of all of the interacting loci must be specified in the regions argument. Any metadata in regions will be preserved.

Note that regions will be resorted in the returned object, so the anchor indices may not be equal to the input anchor1 and anchor2.

Alternatively, GInteractions objects can be constructed by directly supplying the GRanges of the interacting loci to the GInteractions function. If regions is not specified, this will be constructed automatically from the two sets of supplied GRanges. If regions is supplied, exact matching will be performed to identify the indices in regions corresponding to the regions in the supplied anchor GRanges. Missing values are not tolerated and will cause an error to be raised. In both cases, any metadata in the input GRanges will be transferred to the mcols of the output GInteractions object.

All constructors will return a GInteractions object containing all of the specified information. Sorting of regions is performed automatically, with re-indexing of all anchor indices to preserve the correct pairings between regions. If `mode="strict"`, a StrictGInteractions object is returned with anchor indices swapped such that anchor1 <= anchor2 for all interactions. If `mode="reverse"`, a ReverseStrictGInteractions object is returned with anchor indices swapped such that anchor1 >= anchor2. If both anchors are missing, the constructor will return an empty GInteractions object.

Author(s)

Aaron Lun, with contributions from Malcolm Perry and Liz Ing-Simmons.

See Also

interaction-access, interaction-subset, interaction-compare, Vector-class
Examples

```r
set.seed(1000)
N <- 30
all.starts <- round(runif(N, 1, 100))
all.ends <- all.starts + round(runif(N, 5, 20))
all.regions <- GRanges(rep(c("chrA", "chrB"), c(N-10, 10)),
                      IRanges(all.starts, all.ends))

Np <- 20
all.anchor1 <- sample(N, Np)
all.anchor2 <- sample(N, Np)
gi <- GInteractions(all.anchor1, all.anchor2, all.regions)

# Equivalent construction:
gi <- GInteractions(all.regions[all.anchor1], all.regions[all.anchor2])
gi <- GInteractions(all.regions[all.anchor1], all.regions[all.anchor2],
                      all.regions)

# Putting in metadata, elementMetadata
temp.gi <- gi
metadata(temp.gi)$name <- "My GI object"
mcols(temp.gi)$score <- runif(Np)

# Strict construction
sgi <- GInteractions(all.regions[all.anchor1], all.regions[all.anchor2],
                      all.regions, mode="strict")
rsgi <- GInteractions(all.regions[all.anchor1], all.regions[all.anchor2],
                      all.regions, mode="reverse")
```

---

**GRanges methods**

<table>
<thead>
<tr>
<th>Description</th>
<th>Equivalents of GRanges methods</th>
</tr>
</thead>
</table>

**Description**

Methods for GInteractions, InteractionSet and ContactMatrix that operate on the internal GenomicRanges.

**Usage**

```r
## S4 method for signature 'GInteractions'
trim(x, use.names=FALSE)
## S4 method for signature 'GInteractions'
width(x)

# Equivalent calls for InteractionSet and ContactMatrix.
```

**Arguments**

- `x`: A GInteractions, InteractionSet or ContactMatrix object.
- `use.names`: See `"trim,GenomicRanges-method"`. 
**Details**

The `trim` method will return an object of the same class as `x` where out-of-bound ranges in `regions(x)` are truncated to fit. This is followed by resorting of the regions and removal of redundant regions. As such, that the length of `regions(x)` and the anchor indices for each interaction may change before and after trimming. However, the number and order of the interactions will not change.

For GInteractions and InteractionSet objects, the `width` method will return a DataFrame with integer fields `anchor1` and `anchor2`. Each row contains the width of the two anchor regions corresponding to each interaction. For ContactMatrix objects, the method will return a list with the same fields (a different format is necessary as the lengths of anchor1 and anchor2 may be different).

**Value**

Depending on the method, an object of the same class as `x`, or information regarding the genomic regions – see Details.

**Author(s)**

Aaron Lun, based on suggestions from Liz Ing-Simmons.

**See Also**

`trim`, `width`

**Examples**

```r
echo=FALSE)
trim(gi)
width(gi)
trim(x)
width(x)
```

---

**Description**

Methods to get and set fields in an InteractionSet or GInteractions object.

**Usage**

```r
anchors(x, type="both", id=FALSE)
regions(x)
seqinfo(x)
show(x)
```
Interaction accessors

```r
## S4 replacement method for signature 'GInteractions'
anchors(x, type="both") <- value
## S4 replacement method for signature 'GInteractions'
regions(x) <- value
## S4 replacement method for signature 'GInteractions'
seqinfo(x) <- value

## S4 replacement method for signature 'GInteractions'
replaceRegions(x) <- value
## S4 replacement method for signature 'GInteractions'
appendRegions(x) <- value
## S4 method for signature 'GInteractions'
reduceRegions(x)

## S4 method for signature 'GInteractions'
names(x)
## S4 replacement method for signature 'GInteractions'
names(x) <- value

## Each above method has an identical equivalent for
## InteractionSet objects (not shown here for brevity).

## S4 replacement method for signature 'StrictGInteractions'
anchors(x, type="both") <- value
## S4 replacement method for signature 'ReverseStrictGInteractions'
anchors(x, type="both") <- value

## S4 method for signature 'GInteractions'
length(x)
## S4 method for signature 'GInteractions'
as.data.frame(x, row.names=NULL, optional=FALSE, ...)
## S4 method for signature 'GInteractions'
x$name
## S4 replacement method for signature 'GInteractions'
x$name <- value

## S4 method for signature 'InteractionSet'
interactions(x)
## S4 replacement method for signature 'InteractionSet'
interactions(x) <- value
## S4 method for signature 'InteractionSet'
mcols(x, use.names=FALSE)
## S4 replacement method for signature 'InteractionSet'
mcols(x) <- value
```

Arguments

- **x**: An InteractionSet or GInteractions object.
- **type**: a string specifying which anchors are to be extracted or replaced.
- **id**: a scalar logical indicating whether indices or GRanges should be returned.
- **name**: a string indicating the field of mcols to be accessed or modified for a GInteractions object.
**Interaction accessors**

value

For anchors<- and type="first" or "second", an integer vector of length equal to the number of rows in x. For type="both", a list of two such vectors must be supplied.

For regions<-, a GRanges object of length equal to that of regions(x). For replaceRegions<-, a GRanges object that is a superset of all entries in regions(x) involved in interactions. For appendRegions<-, a GRanges of any length containing additional regions.

For seqinfo<-, a SeqInfo object containing chromosome length data for all regions. For interactions<-, a GInteractions object of length equal to that of interactions(x). For mcols<-, a DataFrame with number of rows equal to the length of x. For names<-, a character vector of length equal to that of x.

row.names, optional, ...

Additional arguments, see ?as.data.frame for more details.

use.names

A logical scalar, see ?mcols for more details.

**Value**

For the getters, values in various slots of x are returned, while for the setters, the slots of x are modified accordingly – see below for more details.

**Anchor manipulations for GInteractions**

The return value of anchors varies depending on type and id. If id=FALSE, a GRanges object will be returned containing the regions specified by the anchor1 or anchor2 slots in x, for type="first" or "second", respectively. If type="both", a GRangesList will be returned with two entries - first and second, containing regions specified by anchor1 and anchor2 respectively.

If id=TRUE, the integer vectors in the anchor1 or anchor2 slots of x are returned directly. A list of length two is returned for type="both", containing both of the vectors. Otherwise, each vector is returned corresponding to the requested value of type.

Replacement in anchors<- can only be performed using anchor indices. If type="both", a list of two integer vectors is required in value, specifying the indices of the interacting regions in regions(x). If type="first" or "second", an integer vector is required to replace the existing values in the anchor1 or anchor2 slot, respectively. If the object is a StrictGInteractions, indices are automatically swapped so that anchor1 >= anchor2 for each interaction. The opposite applies if the object is a ReverseStrictGInteractions.

**Region manipulations for GInteractions**

For regions, a GRanges is returned equal to the regions slot in x. For regions<-, the GRanges in value is used to replace the regions slot. Resorting of the replacement GRanges is performed automatically, along with re-indexing of the anchors. In addition, the replacement must be of the same length as the existing object in the regions slot.

The replaceRegions replacement method can take variable length GRanges, but requires that the replacement contain (at least) all ranges contained in anchors(x). The appendRegions replacement method appends extra intervals to the existing regions slot of x. This is more efficient than calling replaceRegions on a concatenated object with regions(x) and the extra intervals.

The reduceRegions method will return a GInteractions object where the regions slot is reduced to only those entries used in anchors(x). This may save some memory in each object by removing unused regions. However, this is not recommended for large workflows with many GInteractions objects. R uses a copy-on-write memory management scheme, so only one copy of the GRanges in regions should be stored so long as it is not modified in different objects.
Interaction accessors

Other methods for GInteractions

For access and setting of all other slots, see Vector-class for details on the appropriate
methods. This includes mcols or metadata, to store interactions-specific or experiment-wide metadata,
respectively. The length method will return the number of interactions stored in x.

The show method will print out the class, the number of pairwise interactions, and the total number
of regions in the GInteractions object. The number and names of the various metadata fields
will also be printed. The as.data.frame method will return a data.frame object containing the
coordinates for the two anchor regions as well as any element-wise metadata.

The seqinfo method will return the sequence information of the GRanges in the regions slot. This
can be replaced with the seqinfo<- method – see ?Seqinfo for more details.

Details for InteractionSet

Almost all InteractionSet methods operate by calling the equivalent method for the GInteractions
object, and returning the resulting value. The only exception is interactions, which returns the
GInteractions object in the interactions slot of the InteractionSet object. This slot can also be set
by supplying a valid GInteractions object in interactions<-. The show method will print information equivalent to that done for a SummarizedExperiment object.

An additional line is added indicating the number of regions in the regions slot of the object.

For access and setting of all other slots, see SummarizedExperiment-class for details on the
appropriate methods. This includes assays, colData, mcols or metadata, which can all be applied
to InteractionSet objects.

Handling different metadata

Note that there are several options for metadata - experiment-wide metadata, which goes into
metadata(x)<-; region-specific metadata (e.g., adjacent genes, promoter/enhancer identity, GC
content), which goes into mcols(regions(x))<-; and interaction-specific metadata (e.g., interac-
tion distance, interaction type), which goes directly into mcols(x)<-. This is applicable to both
GInteractions and InteractionSet objects. In addition, library-specific data (e.g., library size) should
be placed into colData(x)<- for InteractionSet objects.

Users should take care with the differences in the $ and $< operators between these two classes.
For GInteractions objects, this will access or modify fields in the mcols slot, i.e., for interaction-
specific metadata. For InteractionSet objects, this will access or modify fields in the colData slot,
i.e., for library-specific data. The difference in behaviour is due to the concept of libraries in the
InteractionSet, which is lacking in the GInteractions class.

For InteractionSet objects, the mcols getter and setter functions operate on the GInteractions object
stored in interactions slot, rather than accessing the elementMetadata slot of the Summarized-
Experiment base class. This makes no difference for practical usage in the vast majority of cases,
except that any metadata stored in this manner is carried over when the GInteractions object is
extracted with interactions(x). Similarly, the names getter and setter will operate the names
of the internal GInteractions object. However, the metadata getter and setter will operate on the
SummarizedExperiment base class, not on the internal GInteractions object.

Author(s)

Aaron Lun

See Also

GInteractions-class, InteractionSet-class, Vector-class, SummarizedExperiment-class
Examples

```r
eexample(GInteractions, echo=FALSE)  # Generate a nice object.
show(gi)

# Accessing anchor ranges or indices:
anchors(gi)
anchors(gi, type="first")
anchors(gi, id=TRUE)

anchors(gi, id=TRUE, type="first")
anchors(gi, id=TRUE, type="second")

# Modifying anchor indices:
nregs <- length(regions(gi))
mod <- list(sample(nregs, length(gi), replace=TRUE),
            sample(nregs, length(gi), replace=TRUE))
anchors(gi) <- mod
anchors(gi, id=TRUE, type="first")
anchors(gi, id=TRUE, type="second")

anchors(gi, type="both") <- mod
anchors(gi, type="first") <- mod[[1]]
anchors(gi, type="first") <- mod[[2]]

# Accessing or modifying regions:
regions(gi)
reduceRegions(gi)
regions(gi)$score <- runif(length(regions(gi)))

new.ranges <- c(regions(gi), resize(regions(gi), fix="center", width=50))
try(regions(gi) <- new.ranges)  # Fails
replaceRegions(gi) <- new.ranges  # Succeeds

length(regions(gi))
appendRegions(gi) <- GRanges("chrA", IRanges(5:10+1000, 1100+5:10), score=runif(6))
length(regions(gi))

seqinfo(gi)
seqinfo(gi) <- Seqinfo(seqnames=c("chrA", "chrB"), seqlengths=c(1000, 2000))

# Accessing or modifying metadata.
gi$score <- runif(length(gi))
mcols(gi)
as.data.frame(gi)

#################
# Same can be done for an InteractionSet object:

eexample(InteractionSet, echo=FALSE)

anchors(iset)
regions(iset)
reduceRegions(iset)
regions(iset)$score <- regions(gi)$score
replaceRegions(iset) <- new.ranges
```
seqinfo(iset)
seqinfo(iset) <- Seqinfo(seqnames=c("chrA", "chrB"), seqlengths=c(1000, 2000))

# Standard SE methods also available:
colData(iset)
metadata(iset)
mcols(iset)

# Note the differences in metadata storage:
metadata(iset)$name <- "metadata for SE0"
metadata(interactions(iset))$name <- "metadata for GI"

iset$lib.size <- runif(ncol(iset))*1e6
colData(iset)
mcols(iset) # untouched by "$" operator

mcols(iset)$whee <- runif(nrow(iset))
mcols(iset)
mcols(interactions(iset)) # preserved

names(iset) <- paste0("X", seq_along(iset))
names(iset)
names(interactions(iset))

---

Interaction compare

**Interaction comparisons**

**Description**

Methods to order, compare and de-duplicate GInteractions or InteractionSet objects, based on the anchor indices.

**Usage**

```r
## S4 method for signature 'GInteractions'
order(..., na.last=TRUE, decreasing=FALSE)

## S4 method for signature 'GInteractions'
sort(x, decreasing=FALSE, ...)

## S4 method for signature 'GInteractions'
duplicated(x, incomparables=FALSE, fromLast=FALSE, ...)

## S4 method for signature 'GInteractions'
unique(x, incomparables=FALSE, fromLast=FALSE, ...)

## S4 method for signature 'GInteractions'
swapAnchors(x, mode=c("order", "reverse", "all"))

## Each of the above methods has an identical equivalent for
## InteractionSet objects (not shown for brevity).

## S4 method for signature 'GInteractions,GInteractions'
```
match(x, table, nomatch=NA_integer_, incomparables=NULL, ...)

# The same call for 'match' is used for all possible combinations of
# GInteractions and InteractionSet objects in 'x' or 'table'.
# For brevity, these will not be listed here.

## S4 method for signature 'GInteractions,GInteractions'
pcompare(x, y)

### Arguments

... For order, one or more InteractionSet or GInteractions objects with the same
    number of rows. Otherwise, ignored in all other methods.

x An InteractionSet or GInteractions object. For pcompare, a GInteractions object
    only.

na.last A logical scalar indicating whether NA values should be ordered last. This should
    not be relevant as anchor indices should be finite.

decreasing A logical scalar indicating whether rows should be sorted in decreasing order.

incomparables A logical scalar. See ?match for a description of this in match. Otherwise, it is
    ignored.

fromLast A logical scalar indicating whether the last entry of a repeated set in x should be
    considered as a non-duplicate.

mode A string indicating what type of swapping should be performed in swapAnchors.

table A GInteractions or InteractionSet object.

nomatch An integer scalar indicating the value to use for unmatched entries.

y A GInteractions object, of the same length as x.

### Value

For sort and unique, a GInteractions or InteractionSet object is returned, depending on the class
of x.

For order and duplicated, an integer vector of permutations or a logical vector of duplicate spec-
ifications is returned, respectively.

### Sorting and ordering

Sorting is performed based on the anchor indices of the GInteraction object. Rows are ordered for
increasing values of the anchor1 slot - if these are equal, ordering is performed with values of the
anchor2 slot. This equates to ordering by the coordinates directly, as the GRanges in the regions
slot is always sorted. Based on this, sort will return a sorted copy of x with permuted rows in
increasing order.

The order method returns an integer vector indicating the permutation to rearrange x in increasing
order. If multiple objects are supplied to order, ordering will be computed using anchor indices
from successive objects. For example, ordering is first performed using anchor indices from the first
object; any rows with the same anchor1 and anchor2 will be split using the second object; and so
on.

For both of these methods, the sorting can be reversed by setting decreasing=TRUE. This will sort
or order for decreasing values of anchor1 and anchor2, rather than for increasing values.
Removing duplicates

The duplicated method returns a logical vector indicating whether the rows of x are duplicated. Duplicated rows are identified on the basis of identical entries in the anchor1 and anchor2 slots. Values in other slots (e.g., in mcols) are ignored. For a set of duplicated rows, the first occurrence in x is marked as the non-duplicate if fromLast=FALSE, and the last entry otherwise.

unique returns a GInteractions object where all duplicate rows have been removed from x. This is equivalent to x[!duplicated(x),], with any additional arguments passed to duplicated as specified.

Matching and comparing

The match function will return an integer vector of length equal to that of x. Each entry of the vector corresponds to an interaction in x and contains the index of the first interaction table with the same anchor regions. Interactions in x without any matches are assigned NA values by default. Note that the regions slot must be identical between x and table.

The pcompare function will return an integer vector of length equal to x and y. This performs parallel comparisons between corresponding entries in two GInteractions objects, based on the values of the anchor indices (anchor1 first, and then anchor2 if anchor1 is tied). Negative, zero and positive values indicate that the corresponding interaction in x is 'lesser', equal or 'greater' than the corresponding interaction in y. Again, the regions must be identical between the two objects.

Swapping anchors

For GInteractions objects, swapAnchors returns a GInteractions object where anchor1 and anchor2 values are swapped. If mode="order", this is done so that all values in the anchor2 slot are not less than values in anchor1 in the returned object. If mode="reverse", all values in anchor1 are not less than all values in anchor2. If mode="all", the anchor indices in x are directly swapped without consideration of ordering.

It is recommended to apply this method before running methods like order or duplicated. This ensures that redundant permutations are coerced into the same format for a valid comparison. In many applications, permutations of pairwise interactions are not of interest, i.e., an interaction between regions 1 and 2 is the same as that between 2 and 1. Application of swapAnchors with mode="order" ensures that all indices are arranged in a comparable manner. Alternatively, users can use a (Reverse)StrictGInteractions object which enforces a consistent arrangement of indices across interactions.

Methods for InteractionSet objects

For all InteractionSet methods, the corresponding method is called on the GInteractions object in the interactions slot of the InteractionSet object. Return values for each InteractionSet method is the same as those for the corresponding GInteractions method - except for sort and unique, which return a row-permuted or row-subsetted InteractionSet, respectively, instead of a GInteractions object; and swapAnchors, which returns an InteractionSet object where the internal GInteractions has its anchor indices swapped around.

Note that no additional information from the InteractionSet (beyond that in interactions) is used for sorting or duplicate marking, i.e., the assay or metadata values for each interaction are not used in distinguishing rows with the same interaction. For this reason, the pcompare method is not implemented for InteractionSet objects. It makes little sense to do a parallel comparison in an InteractionSet without examining the data.
**Interaction distances**

**Author(s)**

Aaron Lun

**See Also**

`GInteractions-class`, `InteractionSet-class`, `match`, `pcompare`

**Examples**

```r
example(GInteractions, echo=FALSE)

anchors(gi, id=TRUE)
anchors(swapAnchors(gi, mode="all"), id=TRUE)

gi <- swapAnchors(gi)

anchors(gi)

gi2 <- sort(gi)

anchors(gi2)

# Can also order them.
o <- order(gi)

o

stopifnot(all.equal(gi[o], gi2))

# Checking duplication.
summary(duplicated(gi))
summary(duplicated(rbind(gi, gi)))

stopifnot(all.equal(gi, unique(rbind(gi, gi))))

# Matching and comparing.
another.gi <- gi[sample(length(gi))]

match(gi, another.gi)

match(gi, another.gi[1:5])

pcompare(gi, another.gi)

# Same can be done for an InteractionSet object:

example(InteractionSet, echo=FALSE)

iset <- swapAnchors(iset)

anchors(iset)

anchors(sort(iset))

order(iset)

summary(duplicated(iset))

unique(iset)
```

Interaction distances  
*Get the linear distance for each interaction*
Description

Compute the distance between interacting regions on the linear genome, for each pairwise interaction contained in a GInteractions or InteractionSet object.

Usage

```r
## S4 method for signature 'GInteractions'
pairdist(x, type="mid")
## S4 method for signature 'InteractionSet'
pairdist(x, type="mid")

## S4 method for signature 'GInteractions'
intrachr(x)
## S4 method for signature 'InteractionSet'
intrachr(x)
```

Arguments

- `x`: A GInteractions or InteractionSet object.
- `type`: A character string specifying the type of distance to compute. Can take values of "mid", "gap", "span", "diag" or "intra".

Details

For each interaction in `x`, the `pairdist` method computes the distance between the two interacting regions. An integer vector is returned, with values computed according to the specified value of `type`:

- "mid": The distance between the midpoints of the two regions (rounded down to the nearest integer) is returned.
- "gap": The length of the gap between the closest points of the two regions is computed - negative lengths are returned for overlapping regions, indicating the length of the overlap.
- "span": The distance between the furthermost points of the two regions is computed.
- "diag": The difference between the anchor indices is returned. This corresponds to a diagonal on the interaction space when bins are used in the `regions` slot of `x`.

Interchromosomal interactions are marked with NA. Alternatively, if `type="intra"`, a logical vector is returned indicating whether the interaction occurs between two regions on the same chromosome. `intrachr(x)` is an alias for `pairdist(x, type="intra")`.

The return values of the assorted methods are the same for both GInteractions and InteractionSet objects. Methods for the latter operate on the GInteractions object in the `interactions` slot.

Value

An integer or logical vector of distances.

Author(s)

Aaron Lun

See Also

`GInteractions-class`, `InteractionSet-class`
Interaction overlaps

Examples

```r
example(GInteractions, echo=FALSE)
pairdist(gi)
pairdist(gi, type="gap")
pairdist(gi, type="span")
pairdist(gi, type="diag")
intrachr(gi)
```

```r
example(InteractionSet, echo=FALSE)
pairdist(iset)
pairdist(iset, type="gap")
pairdist(iset, type="span")
pairdist(iset, type="diag")
intrachr(iset)
```

Interaction overlaps  Find overlaps between interactions in one or two dimensions

Description

Find overlaps between interactions and linear intervals, between interactions and pairs of intervals, and between interactions and other interactions in a GInteractions or InteractionSet object.

Usage

```r
## S4 method for signature 'GInteractions,Vector'
findOverlaps(query, subject, maxgap=0L, minoverlap=1L,
type=c("any", "start", "end", "within", "equal"),
select=c("all", "first", "last", "arbitrary"),
ignore.strand=FALSE, use.region="both")
```

```r
## S4 method for signature 'GInteractions,Vector'
overlapsAny(query, subject, maxgap=0L, minoverlap=1L,
type=c("any", "start", "end", "within", "equal"),
ignore.strand=FALSE, use.region="both")
```

```r
## S4 method for signature 'GInteractions,Vector'
countOverlaps(query, subject, maxgap=0L, minoverlap=1L,
type=c("any", "start", "end", "within", "equal"),
ignore.strand=FALSE, use.region="both")
```

```r
## S4 method for signature 'GInteractions,Vector'
subsetByOverlaps(query, subject, maxgap=0L, minoverlap=1L,
type=c("any", "start", "end", "within", "equal"),
ignore.strand=FALSE, use.region="both")
```

# For brevity, only 'GInteractions,Vector-methods' are listed. Methods are available for all pairwise combinations of GInteractions, InteractionSet, and Vector objects, so long as at least one InteractionSet or GInteractions object is present. In all cases, function calls are identical. 'subject' can
# also be missing for all functions except for 'subsetByOverlaps', as long as
# 'query' is a GInteractions or an InteractionSet.

## Arguments

query, subject  
A Vector, GInteractions or InteractionSet object, depending on the specified method. At least one of these must be a GInteractions or InteractionSet object. Also, subject can be missing if query is a GInteractions or InteractionSet object.

maxgap, minoverlap, type  
See ?findOverlaps in the GenomicRanges package.

select, ignore.strand  
See ?findOverlaps in the GenomicRanges package.

use.region  
A string specifying the regions to be used to identify overlaps.

## Value

For findOverlaps, a Hits object is returned if select="all", and an integer vector of subject indices otherwise.

For countOverlaps and overlapsAny, an integer or logical vector is returned, respectively.

For subsetByOverlaps, a subsetted object of the same class as query is returned.

### Overview of overlaps for GInteractions

For all methods taking a Vector and an GInteractions, the Vector is assumed to represent some region on the linear genome (e.g., GRanges) or set of such regions (GRangesList). An overlap will be defined between the interval and an GInteractions interaction if either anchor region of the latter overlaps the former. This is considered to be a one-dimensional overlap, i.e., on the linear genome.

For methods between two GInteractions objects, a two-dimensional overlap will be computed between the anchor regions of the two objects. An overlap is defined if each anchor region of the first object overlaps at least one anchor region of the second object, and each anchor region of the second object overlaps at least one anchor region of the first object, i.e., there are overlapping areas in the two-dimensional interaction space. If subject is missing, overlaps will be computed between interactions in query.

### Description of overlap methods

When select="all", findOverlaps returns a Hits object containing overlapping pairs of queries and subjects (or more specifically, their indices in the supplied objects - see ?findOverlaps for more details). For other values of select, an integer vector is returned with one entry for each element of query, which specifies the index of the chosen (first, last or arbitrary) overlapping feature in subject for that query. Queries with no overlaps at all are assigned NA values.

For the other methods, countOverlaps returns an integer vector indicating the number of elements in subject that were overlapped by each element in query. overlapsAny returns a logical vector indicating which elements in query were overlapped by at least one element in subject. subsetByOverlaps returns a subsetted query containing only those elements overlapped by at least one element in subject.
Interaction overlaps

Choice of regions to define overlaps

For one-dimensional overlaps, use.region="both" by default such that overlaps with either interacting region are considered. If use.region="first", overlaps are only considered between the interval and the first interacting region. Similarly, if use.region="second", only the second interaction region is used.

For two-dimensional overlaps, use.region="both" by default such that the order of first/second interacting/target regions is ignored. This means that an overlap will be considered between, e.g., the first interacting region and the second target region. If use.region="same", overlaps are only considered between the set of first regions for the GInteractions object and the first regions for the target pairs, and similarly for the second regions. If use.region="reverse", overlaps are only considered between the set of first regions for the GInteractions object and the second regions for the target pairs, and vice versa. These options tend only to be useful if the order of first/second regions is informative.

Details for InteractionSet

The behaviour of each method for InteractionSet objects is largely the same as that described for GInteractions objects. For a given InteractionSet object x, the corresponding method is called on the GInteractions object in the interactions slot of x. The return value is identical to that from calling the method on interactions(x), except for subsetByOverlaps for InteractionSet queries (which returns a subsetted InteractionSet object, containing only those rows/interactions overlapping the subject).

Author(s)

Aaron Lun

See Also

InteractionSet-class, findOverlaps, linkOverlaps

Examples

element(GInteractions, echo=FALSE)

# Making a larger object, for more overlaps.
Np <- 100
N <- length(regions(gi))
all.anchor1 <- sample(N, Np, replace=TRUE)
all.anchor2 <- sample(N, Np, replace=TRUE)
gi <- GInteractions(all.anchor1, all.anchor2, regions(gi))

# GRanges overlaps:
of.interest <- resize(sample(regions(gi), 2), width=1, fix="center")
findOverlaps(of.interest, gi)
findOverlaps(gi, of.interest)
findOverlaps(gi, of.interest, select="first")
overlapsAny(gi, of.interest)
overlapsAny(of.interest, gi)
countOverlaps(gi, of.interest)
countOverlaps(of.interest, gi)
subsetByOverlaps(gi, of.interest)
subsetByOverlaps(of.interest, gi)
Interaction subsetting

Methods to subset or combine InteractionSet or GIinteractions objects.
## Interaction subsetting

### Usage

```r
## S4 method for signature 'GInteractions,ANY,ANY'
x[i, j, ..., drop=TRUE]
## S4 replacement method for signature 'GInteractions,ANY,ANY,GInteractions'
x[i, j] <- value

## S4 method for signature 'GInteractions'
rbind(..., deparse.level=1)
## S4 method for signature 'GInteractions'
c(x, ..., recursive=FALSE)

## S4 method for signature 'GInteractions'
split(x, f, drop=FALSE)

## Each of the above methods has an identical equivalent for
## InteractionSet objects (not shown for brevity).

## S4 method for signature 'InteractionSet'
subset(x, i, j)
## S4 method for signature 'InteractionSet'
cbind(..., deparse.level=1)
```

### Arguments

- **x**: A GInteractions or InteractionSet object.
- **i, j**: A vector of logical or integer subscripts. For InteractionSet objects, these indicate the rows and columns to be subsetted for i and j, respectively. Rows correspond to pairwise interactions while columns correspond to samples. For GInteractions objects, i indicates the genomic interactions to be retained. j is ignored as there is no concept of samples in this class.
- **value**: A GInteractions or InteractionSet object with length or number of rows equal to length of i (or that of x, if i is not specified). For InteractionSet objects, the number of columns must be equal to the length of j (or number of columns in x, if j is not specified).
- **...**: For rbind and c, ... contains GInteractions or InteractionSet objects to be combined row-wise, depending on the method. All objects must be of the same class. For c, any objects are additional to that already specified in x. For cbind, ... contains InteractionSet objects to be combined column-wise. For all other methods, this argument is ignored.
- **f**: A vector-like object of length equal to length(x) (for GInteractions objects) or nrow(x) (for InteractionSet objects), indicating how entries/rows of x are to be split into separate objects. This is usually a factor.
- **drop**: A logical scalar, indicating if levels of f that do not occur should be dropped in split, InteractionSet-method. This is ignored by the other methods.
- **deparse.level**: An integer scalar; see ?base::cbind for a description of this argument.
- **recursive**: An integer scalar, ignored.

### Value

A subsetted or combined object of the same class as x.
Details for GInteractions

Subsetting operations will return a GInteractions object containing the specified interactions. Values of the anchor1 and anchor2 slots will be appropriately subsetted in the returned object, along with any metadata in mcols. However, note that the value of regions will not be modified by subsetting. rbind and c are synonyms and will concatenate GInteractions objects. These methods will check whether the regions slot of all supplied objects are the same, in which case the regions and anchor indices are used directly. Otherwise, a new GRanges is constructed containing the (sorted) union of all regions across the input objects. Anchor indices in each object are refactored appropriately to refer to the relevant entries in this new object.

Note that the column names in mcols must be identical across all supplied objects in rbind and c. The column names of mcols for the regions slot must also be identical across objects. If GInteractions objects of different strictness (i.e., StrictGInteractions and ReverseGInteractions) are concatenated, the returned object will be of the same class as the first supplied object.

split produces a list of GInteractions objects, where each entry of the list corresponds to a level of f. Each GInteractions object contains all rows associated with the corresponding level of f.

For short index vectors, subsetting a GInteractions object prior to calling anchors may be much faster than the reverse procedure. This is because the anchors getter will construct a GRanges(List) containing the genomic loci for all pairwise interactions. Subsetting beforehand ensures that only loci for the desired interactions are included. This avoids constructing the entire object just to subset it later.

Details for InteractionSet

Subsetting behaves in much the same way as that for the SummarizedExperiment class. Interactions are treated as rows and will be subsetted as such. All subsetting operations will return an InteractionSet with the specified interactions (rows) or samples (columns). Again, note that the value of regions will not be modified by subsetting.

cbind will combine objects with the same interactions but different samples. Errors will occur if the interactions are not identical between objects (i.e., must have same values in the interactions slots). Additional restrictions apply on the column and assay names - see cbind,SummarizedExperiment-method for details.

rbind and c will combine objects with the same samples but different interactions. In this case, the interactions need not be identical, and will be combined using the methods described above for GInteractions objects. Again, additional restrictions apply - see rbind,SummarizedExperiment-method for details.

split produces a list of InteractionSet objects, where each entry of the list corresponds to a level of f. Each InteractionSet object contains all rows associated with the corresponding level of f.

Author(s)

Aaron Lun

See Also

InteractionSet-class

Examples

example(GInteractions, echo=FALSE)

# Subsetting:
InteractionSet class

```r
gi[1,]
gi[1:2,]
gi[3]
gi[3:4]

temp.gi <- gi

# Combining:
rbindi(gi, gi)
c(gi, gi)

new.gi <- gi
regions(new.gi) <- resize(regions(new.gi), width=20, fix="start")
rbindi(gi, new.gi)

# Splitting:
f <- sample(4, length(gi), replace=TRUE)
out <- split(gi, f)
out[[1]]

# Same can be done for an InteractionSet object:

eexample(InteractionSet, echo=FALSE)

# Subsetting:
iset[1,]
iset[1:2,]
iset[1,1]
iset[1:2,1]
iset[3,3]
iset[3:4,3:4]

# Combining:
cbindi(iset, iset)
rbindi(iset, iset)
c(iset, iset)

new.iset <- iset
regions(new.iset) <- resize(regions(new.iset), width=20, fix="start")
rbindi(iset, new.iset)

# Splitting:
out <- split(iset, f)
out[[1]]
```

### Description

The InteractionSet class stores information about pairwise genomic interactions, and is intended for use in data analysis from Hi-C or ChIA-PET experiments. Each row of the InteractionSet corresponds to a pairwise interaction between two loci, as defined in the GInteractions object. Each
column corresponds to a library or sample. Each InteractionSet also contains one or more assays, intended to hold experimental data about interaction frequencies for each interaction in each sample.

Usage

```r
## S4 method for signature 'ANY,GInteractions'
InteractionSet(assays, interactions, ...)
```

```r
## S4 method for signature 'missing,missing'
InteractionSet(assays, interactions, ...)
```

Arguments

- **assays**: A numeric matrix or a list or `SimpleList` of matrices, containing data for each interaction.
- **interactions**: A GInteractions object of length equal to the number of rows in `assays`.
- **...**: Other arguments to be passed to `SummarizedExperiment`.

Details

The InteractionSet class inherits from the SummarizedExperiment class and has access to all of its data members and methods (see `?SummarizedExperiment-class` for more details). It also contains an additional interactions slot which holds a GInteractions object (or an object from any derived classes, e.g., StrictGInteractions). Each row of the InteractionSet object corresponds to a pairwise interaction between two genomic loci in `interactions`.

The constructor will return an InteractionSet object containing all of the specified information - for `InteractionSet,missing,missing-method`, an empty InteractionSet object is returned. Note that any metadata arguments will be placed in the metadata of the internal SummarizedExperiment object, not the metadata of the internal GInteractions object. This is consistent with the behaviour of similar classes like RangedSummarizedExperiment.

Value

For the constructors, an InteractionSet object is returned.

Author(s)

Aaron Lun

See Also

`SummarizedExperiment, interaction-access, interaction-subset, interaction-compare, SummarizedExperiment-class`

Examples

```r
example(GInteractions, echo=FALSE)
Nlibs <- 4
counts <- matrix(rpois(Np*Nlibs, lambda=10), ncol=Nlibs)
colnames(counts) <- seq_len(Nlibs)
iset <- InteractionSet(counts, gi)
iset <- InteractionSet(counts, gi, colData=DataFrame(lib.size=1:Nlibs*1000))
iset <- InteractionSet(counts, gi, metadata=list(name="My Hi-C data"))
```
Note differences in metadata storage:
metadata(iset)
metadata(interactions(iset))

Linearize interactions

**Description**

Convert interactions in two-dimensional space to one-dimensional ranges on the linear genome.

**Usage**

```r
## S4 method for signature 'GInteractions,numeric'
linearize(x, ref, internal=TRUE)

## S4 method for signature 'GInteractions,GRanges'
linearize(x, ref, ..., internal=TRUE)

# Equivalent calls for InteractionSet objects.
```

**Arguments**

- `x` A GInteractions or InteractionSet object
- `ref` A numeric vector or a GRanges object, specifying the reference region(s) to use for linearization. If numeric, the entries should be indices pointing to a genomic interval in `regions(x)`.
- `internal` A logical scalar specifying whether interactions within `ref` should be reported.
- `...` Other arguments, passed to `overlapsAny` in the GRanges methods.

**Details**

This method identifies all interactions with at least one anchor region overlapping the specified region(s) in `ref`. When `x` is a GInteractions object, the method returns a GRanges object with one entry per identified interaction, where the coordinates are defined as the *other* anchor region, i.e., the one that does not overlap the reference region.

If both of the anchor regions for an interaction overlap the reference regions, the genomic interval spanned by both anchor regions is returned. This is because it is not clear which region should be defined as the "other" anchor in such circumstances. Note that this will fail if the reference regions occur across multiple chromosomes. If `internal=FALSE`, interactions with both overlapping anchor regions are removed from the output.

When `x` is an InteractionSet object, a RangedSummarizedExperiment object is returned. Each entry corresponds to an identified interaction with the non-overlapping anchor region stored in the `rowRanges`. Experimental data associated with each identified interaction is stored in the various assays.

This method effectively converts two-dimensional interaction data into one-dimensional coverage across the linear genome. It is useful when a particular genomic region is of interest - this can be
used as ref, to examine the behaviour of all other regions relative to it. For example, Hi-C data in x can be converted into pseudo-4C contact frequencies after linearization. Disjoint ranges across multiple chromosomes are supported when ref is a GRanges object. However, it usually only makes sense to use contiguous ranges as a single bait region. Similarly, if ref is numeric, it should refer to consecutive entries in regions(x) to specify the bait region.

Value

A GRanges when x is a GInteractions object, and a RangedSummarizedExperiment when x is an InteractionSet object.

Examples

example(InteractionSet, echo=FALSE)

# With integers
out <- linearize(iset, 1)
linearize(iset, 10)
linearize(iset, 20)

# With ranges
linearize(iset, regions(iset)[1], type="equal")
linearize(iset, regions(iset)[10], type="equal")
linearize(iset, regions(iset)[20], type="equal")
**Details**

This function identifies all interactions in `query` where one anchor overlaps an entry in `subject1` and the other anchor overlaps an entry in `subject2`. It is designed to be used to identify regions that are linked by interactions in `query`. For example, one might specify genes as `subject1` and enhancers as `subject2`, to identify all gene-enhancer contacts present in `query`. This is useful when the exact pairings between `subject1` and `subject2` are undefined.

The function returns a dataframe specifying the index of the interaction in `query`; the index of the overlapped region in `subject1`; and the index of the overlapped region in `subject2`. If multiple regions in `subject1` and/or `subject2` are overlapping the anchor regions of a particular interaction, all combinations of two overlapping regions (one from each `subject*` set) are reported for that interaction.

By default, `use.region="both"` such that overlaps will be considered between any first/second interacting region in `query` and either `subject1` or `subject2`. If `use.region="same"`, overlaps will only be considered between the first interacting region in `query` and entries in `subject1`, and between the second interacting region and `subject2`. The opposite applies with `use.region="reverse"`, where the first and second interacting regions are overlapped with `subject2` and `subject1` respectively.

If `subject2` is not specified, links within `subject1` are identified instead, i.e., `subject2` is set to `subject1`. In such cases, the returned dataframe is such that the first subject index is always greater than the second subject index, to avoid redundant permutations.

**Value**

A dataframe of integer indices indicating which elements of `query` link which elements of `subject1` and `subject2`.

**Author(s)**

Aaron Lun

**See Also**

`findOverlaps`, `GInteractions`, `Vector-method`

**Examples**

```r
eexample(GInteractions, echo=FALSE)

all.genes <- GRanges("chrA", IRanges(0:9*10, 1:10*10))
all.enhancers <- GRanges("chrB", IRanges(0:9*10, 1:10*10))

out <- linkOverlaps(gi, all.genes, all.enhancers)
head(out)

out <- linkOverlaps(gi, all.genes)
head(out)

# Same methods apply for InteractionSet objects.

eexample(InteractionSet, echo=FALSE)
out <- linkOverlaps(iset, all.genes, all.enhancers)
out <- linkOverlaps(iset, all.genes)
```
pairs

## Extract paired ranges

**Description**

Represent interactions in a GInteractions or Interaction object as a Pairs, SelfHits or GRangesList object.

**Usage**

```
## S4 method for signature 'GInteractions'
pairs(x, id=FALSE, as.grlist=FALSE)
```

# Equivalent call for InteractionSet to above.

```
makeGInteractionsFromGRangesPairs(x)
```

**Arguments**

- **x**
  - For `pairs`, a GInteractions or InteractionSet object. For `makeGInteractionsFromGRangesPairs`, a Pairs object containing two parallel GRanges.
- **id**
  - A logical scalar specifying whether indices should be returned instead of regions.
- **as.grlist**
  - A logical scalar specifying whether a GRangesList should be returned.

**Details**

Recall that the GInteractions object stores anchor regions for each interaction in two parallel GRanges, where corresponding entries between the two GRanges constitute the pair of regions for one interaction. These parallel ranges can be extracted and stored as a Pairs object for further manipulation. This is similar to the GRangesList reported by `anchors` with `type="both"` and `id=FALSE`. The reverse conversion is performed using `makeGInteractionsFromGRangesPairs`.

An alternative representation involves storing the two anchors for each interaction in a single GRanges of length 2. Multiple interactions are then stored as a GRangesList, along with any metadata and sequence information. This is returned if `as.grlist=FALSE`, may be more useful in some applications where the two interacting regions must be in the same GRanges. Finally, if `id=TRUE`, the anchor indices are extracted and returned as a SelfHits object. This may be useful for graph construction.

**Value**

- For `pairs`, if `id=TRUE`, a SelfHits object is returned. Otherwise, if `as.grlist=TRUE`, a GRangesList object is returned. Otherwise, a Pairs object is returned.
- For `makeGInteractionsFromGRangesPairs`, a GInteractions object is returned.

**Author(s)**

Aaron Lun
See Also

`GInteractions, Pairs, SelfHits, GRangesList`

Examples

```r
e.example(GInteractions, echo=FALSE)
y <- pairs(gi)
y
makeGInteractionsFromGRangesPairs(y)

pairs(gi, id=TRUE)
pairs(gi, as.grlist=TRUE)

e.example(InteractionSet, echo=FALSE)
pairs(iset)
pairs(iset, id=TRUE)
pairs(iset, as.grlist=TRUE)
```
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