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Contents

1	Introduction		
2	GRanges: Genomic Ranges		2
	2.1	Splitting and combining <i>GRanges</i> objects	4
	2.2	Subsetting GRanges objects	5
	2.3	Basic interval operations for <i>GRanges</i> objects	8
	2.4	Interval set operations for <i>GRanges</i> objects	11
3	GRangesList: Groups of Genomic Ranges		
	3.1	Basic GRangesList accessors	13
	3.2	Combining GRangesList objects	15
	3.3	Basic interval operations for <i>GRangesList</i> objects	17
	3.4	Subsetting GRangesList objects	18
	3.5	Looping over GRangesList objects	20
4	Interval overlaps involving GRanges and GRangesList objects		23
5	Session Information		

1 Introduction

The *GenomicRanges* package serves as the foundation for representing genomic locations within the *Bioconductor* project. In the *Bioconductor* package hierarchy, it builds upon the *IRanges* (infrastructure) package and provides support for the *BSgenome* (infrastructure), *Rsamtools* (I/O), *ShortRead* (I/O & QA), *rtracklayer* (I/O), *GenomicFeatures* (infrastructure), *GenomicAlignments* (sequence reads), *VariantAnnotation* (called variants), and many other *Bioconductor* packages.

This package lays a foundation for genomic analysis by introducing three classes (*GRanges*, *GPos*, and *GRangesList*), which are used to represent genomic ranges, genomic positions, and groups of genomic ranges. This vignette focuses on the *GRanges* and *GRangesList* classes and their associated methods.

The *GenomicRanges* package is available at https://bioconductor.org and can be installed via biocLite:

```
> source("https://bioconductor.org/biocLite.R")
> biocLite("GenomicRanges")
```

A package only needs to be installed once. Load the package into an R session with

> library(GenomicRanges)

2 GRanges: Genomic Ranges

The *GRanges* class represents a collection of genomic ranges that each have a single start and end location on the genome. It can be used to store the location of genomic features such as contiguous binding sites, transcripts, and exons. These objects can be created by using the *GRanges* constructor function. For example,

```
> gr <- GRanges(
      seqnames = Rle(c("chr1", "chr2", "chr1", "chr3"), c(1, 3, 2, 4)),
      ranges = IRanges(101:110, end = 111:120, names = head(letters, 10)),
      strand = Rle(strand(c("-", "+", "*", "+", "-")), c(1, 2, 2, 3, 2)),
      score = 1:10,
      GC = seq(1, 0, length=10))
> gr
GRanges object with 10 ranges and 2 metadata columns:
    segnames
                ranges strand |
                                                          GC
                                     score
       <Rle> <IRanges> <Rle> | <integer>
                                                   <numeric>
       chr1 [101, 111]
                             - |
  а
       chr2 [102, 112]
                                         2 0.88888888888889
  b
                             + |
  C
       chr2 [103, 113]
                            + |
                                         3 0.7777777777778
                           . . . .
       chr3 [108, 118]
                                         8 0.2222222222222
  h
                            + |
        chr3 [109, 119]
                                         9 0.111111111111111
        chr3 [110, 120]
                             - |
                                        10
  i
  seqinfo: 3 sequences from an unspecified genome; no seqlengths
> options(warn=2)
```

creates a *GRanges* object with 10 genomic ranges. The output of the *GRanges* show method separates the information into a left and right hand region that are separated by | symbols. The genomic coordinates (seqnames, ranges, and strand) are located on the left-hand side and the metadata columns (annotation) are located on the right. For this example, the metadata is comprised of score and GC information, but almost anything can be stored in the metadata portion of a *GRanges* object.

The components of the genomic coordinates within a *GRanges* object can be extracted using the seqnames, ranges, and strand accessor functions.

```
> seqnames(gr)
```

```
factor-Rle of length 10 with 4 runs
            1 3 2
  Lengths:
  Values : chr1 chr2 chr1 chr3
Levels(3): chr1 chr2 chr3
> ranges(gr)
IRanges object with 10 ranges and 0 metadata columns:
       start
                 end
                          width
   <integer> <integer> <integer>
         101
                  111
         102
                  112
                             11
 h
  С
         103
                 113
                           11
         . . .
                  . . .
         108
                  118
                            11
  h
        109
 i
                  119
                           11
 j
         110
                  120
                            11
> strand(gr)
factor-Rle of length 10 with 5 runs
  Lengths: 1 2 2 3 2
  Values : - + * + -
Levels(3): + - *
```

The genomic ranges can be extracted without corresponding metadata with granges

```
> granges(gr)
GRanges object with 10 ranges and 0 metadata columns:
    segnames
               ranges strand
      <Rle> <IRanges> <Rle>
    chr1 [101, 111]
  b
     chr2 [102, 112]
      chr2 [103, 113]
        . . .
                   . . .
       chr3 [108, 118]
      chr3 [109, 119]
  i
       chr3 [110, 120]
  j
  seqinfo: 3 sequences from an unspecified genome; no seqlengths
```

Annotations for these coordinates can be extracted as a *DataFrame* object using the mcols accessor.

```
8 8 0.2222222

9 9 0.111111

10 10 0.0000000

> mcols(gr)$score

[1] 1 2 3 4 5 6 7 8 9 10
```

Information about the lengths of the various sequences that the ranges are aligned to can also be stored in the *GRanges* object. So if this is data from *Homo sapiens*, we can set the values as:

```
> seqlengths(gr) <- c(249250621, 243199373, 198022430)
```

And then retrieves as:

```
> seqlengths(gr)
chr1 chr2 chr3
249250621 243199373 198022430
```

Methods for accessing the length and names have also been defined.

```
> names(gr)
[1] "a" "b" "c" "d" "e" "f" "g" "h" "i" "j"
> length(gr)
[1] 10
```

2.1 Splitting and combining *GRanges* objects

GRanges objects can be devided into groups using the split method. This produces a GRangesList object, a class that will be discussed in detail in the next section.

```
> sp <- split(gr, rep(1:2, each=5))</pre>
> sp
GRangesList object of length 2:
GRanges object with 5 ranges and 2 metadata columns:
                                                        GC
   segnames
                ranges strand |
                                   score
      <Rle> <IRanges> <Rle> | <integer>
                                                 <numeric>
       chrl [101, 111]
                           - |
 а
                           + |
                                       2 0.88888888888889
 b
       chr2 [102, 112]
                                     3 0.7777777777778
       chr2 [103, 113]
                          + |
 С
       chr2 [104, 114]
                          *
                                     4 0.66666666666667
       chr1 [105, 115]
                                       5 0.555555555556
                           *
 е
$2
GRanges object with 5 ranges and 2 metadata columns:
   segnames
              ranges strand | score
                                                    GC
       chr1 [106, 116]
                       + | 6 0.444444444444444
```

Separate *GRanges* instances can be concatenated by using the c and append methods.

```
> c(sp[[1]], sp[[2]])
GRanges object with 10 ranges and 2 metadata columns:
                ranges strand |
                                     score
                                                            GC
       <Rle> <IRanges> <Rle> | <integer>
                                                     <numeric>
       chr1 [101, 111]
                           - | 1
  а
        chr2 [102, 112]
                              + |
                                          2 0.88888888888889
  b
       chr2 [103, 113]
                            + |
                                        3 0.7777777777778
                           . . . .
                                       . . .
       chr3 [108, 118] + |
chr3 [109, 119] - |
chr3 [110, 120] - |
                                       8 0.2222222222222
9 0.11111111111111
                                        10
  j
  seqinfo: 3 sequences from an unspecified genome
```

2.2 Subsetting GRanges objects

GRanges objects act like vectors of ranges, with the expected vector-like subsetting operations available

A second argument to the [subset operator can be used to specify which metadata columns to extract from the *GRanges* object. For example,

Elements can also be assigned to the *GRanges* object. Here is an example where the second row of a *GRanges* object is replaced with the first row of gr.

```
> singles <- split(gr, names(gr))</pre>
> grMod <- gr
> grMod[2] <- singles[[1]]
> head(grMod, n=3)
GRanges object with 3 ranges and 2 metadata columns:
    seanames
               ranges strand |
                                  score
                                                       GC
      <Rle> <IRanges> <Rle> | <integer>
                                                <numeric>
       chr1 [101, 111]
                         - | 1
                                                        1
                           - |
       chr1 [101, 111]
                                                        1
       chr2 [103, 113]
                          + |
                                     3 0.7777777777778
  seqinfo: 3 sequences from an unspecified genome
```

Here is a second example where the metadata for score from the third element is replaced with the score from the second row etc.

```
> grMod[2,1] <- singles[[3]][,1]
> head(grMod, n=3)
GRanges object with 3 ranges and 2 metadata columns:
   segnames
              ranges strand |
                                                     GC
                                 score
      <Rle> <IRanges> <Rle> | <integer>
                                               <numeric>
       chr1 [101, 111] - | 1
                                                       1
 а
       chr2 [103, 113]
 b
                          + |
                                                       1
       chr2 [103, 113]
                         + |
                                    3 0.7777777777778
 seqinfo: 3 sequences from an unspecified genome
```

There are methods to repeat, reverse, or select specific portions of GRanges objects.

```
> rep(singles[[2]], times = 3)
GRanges object with 3 ranges and 2 metadata columns:
   segnames
             ranges strand |
                               score
                                                 GC
     <Rle> <IRanges> <Rle> | <integer>
                                           <numeric>
      chr2 [102, 112]
                        + |
                                 2 0.88888888888888
      chr2 [102, 112]
                    + |
                                 2 0.88888888888888
 seqinfo: 3 sequences from an unspecified genome
> rev(gr)
GRanges object with 10 ranges and 2 metadata columns:
                                                 GC
   segnames
             ranges strand |
                              score
     <Rle> <IRanges> <Rle> | <integer>
                               10
 j
      chr3 [110, 120]
                      - |
                                9 0.11111111111111
      chr3 [109, 119]
 i
                        - |
                                 8 0.2222222222222
      chr3 [108, 118]
                       + |
```

```
. ... ... ... ...
   chr2 [103, 113] + | 3 0.777777777778
chr2 [102, 112] + | 2 0.8888888888888888889
chr1 [101, 111] - | 1
 h
 a chr1 [101, 111] - |
 seginfo: 3 sequences from an unspecified genome
> head(gr,n=2)
GRanges object with 2 ranges and 2 metadata columns:
   seqnames ranges strand | score
 seqinfo: 3 sequences from an unspecified genome
> tail(gr,n=2)
GRanges object with 2 ranges and 2 metadata columns:
   seqnames ranges strand | score
    <Rle> <IRanges> <Rle> | <integer> <numeric>
     chr3 [109, 119] - | 9 0.11111111111111
                                 10
 j
     chr3 [110, 120]
                       - |
 seqinfo: 3 sequences from an unspecified genome
> window(gr, start=2,end=4)
GRanges object with 3 ranges and 2 metadata columns:
            ranges strand | score
   segnames
                                                 GC
    <Rle> <IRanges> <Rle> | <integer> <numeric>
 b chr2 [102, 112] + | 2 0.888888888888889
c chr2 [103, 113] + | 3 0.7777777777778
d chr2 [104, 114] * | 4 0.6666666666666667
 seqinfo: 3 sequences from an unspecified genome
> gr[IRanges(start=c(2,7), end=c(3,9))]
GRanges object with 5 ranges and 2 metadata columns:
   segnames ranges strand | score
     <Rle> <IRanges> <Rle> | <integer> <numeric>
 seqinfo: 3 sequences from an unspecified genome
```

2.3 Basic interval operations for *GRanges* objects

Basic interval characteristics of *GRanges* objects can be extracted using the start, end, width, and range methods.

```
> q <- qr[1:3]
> g <- append(g, singles[[10]])</pre>
> start(g)
[1] 101 102 103 110
> end(g)
[1] 111 112 113 120
> width(q)
[1] 11 11 11 11
> range(q)
GRanges object with 3 ranges and 0 metadata columns:
      segnames
                   ranges strand
         <Rle> <IRanges> <Rle>
  [1]
          chr1 [101, 111]
          chr2 [102, 113]
  [2]
  [3]
          chr3 [110, 120]
  seqinfo: 3 sequences from an unspecified genome
```

The *GRanges* class also has many methods for manipulating the ranges. The methods can be classified as *intra-range methods*, *inter-range methods*, and *between-range methods*.

Intra-range methods operate on each element of a GRanges object independent of the other ranges in the object. For example, the flank method can be used to recover regions flanking the set of ranges represented by the GRanges object. So to get a GRanges object containing the ranges that include the 10 bases upstream of the ranges:

```
> flank(g, 10)
GRanges object with 4 ranges and 2 metadata columns:
   segnames
                ranges strand |
                                                       GC
      <Rle> <IRanges> <Rle> | <integer>
                                                <numeric>
       chr1 [112, 121]
                         - | 1
       chr2 [ 92, 101]
                           + |
                                       2 0.88888888888889
       chr2 [ 93, 102]
                           + |
                                      3 0.7777777777778
 c
       chr3 [121, 130]
                          - |
  seqinfo: 3 sequences from an unspecified genome
```

And to include the downstream bases:

```
> flank(g, 10, start=FALSE)

GRanges object with 4 ranges and 2 metadata columns:
    seqnames ranges strand | score GC
    <Rle> <IRanges> <Rle> | <integer> <numeric>
```

Other examples of intra-range methods include resize and shift. The shift method will move the ranges by a specific number of base pairs, and the resize method will extend the ranges by a specified width.

```
> shift(g, 5)
GRanges object with 4 ranges and 2 metadata columns:
   segnames
               ranges strand |
                                   score
                                                       GC
      <Rle> <IRanges> <Rle> | <integer>
                                                <numeric>
       chr1 [106, 116]
                       - |
                                  1
       chr2 [107, 117]
                          + |
                                      2 0.8888888888888
       chr2 [108, 118]
                          + |
                                     3 0.7777777777778
       chr3 [115, 125]
                           - |
                                    10
                                                        0
 seqinfo: 3 sequences from an unspecified genome
> resize(g, 30)
GRanges object with 4 ranges and 2 metadata columns:
                                                       GC
   segnames
               ranges strand |
                                   score
      <Rle> <IRanges> <Rle> | <integer>
                                                <numeric>
       chr1 [ 82, 111]
                          - |
                                                        1
 а
       chr2 [102, 131]
                           + |
                                      2 0.88888888888889
                                     3 0.7777777777778
       chr2 [103, 132]
                           + |
 С
       chr3 [ 91, 120]
                                     10
 j
                          - |
 seqinfo: 3 sequences from an unspecified genome
```

The GenomicRanges help page ?"intra-range-methods" summarizes these methods.

Inter-range methods involve comparisons between ranges in a single *GRanges* object. For instance, the <u>reduce</u> method will align the ranges and merge overlapping ranges to produce a simplified set.

Sometimes one is interested in the gaps or the qualities of the gaps between the ranges represented by your *GRanges* object. The gaps method provides this information: reduced version of your ranges:

```
> gaps(g)
GRanges object with 12 ranges and 0 metadata columns:
                          ranges strand
       segnames
          <Rle>
                       <IRanges> <Rle>
   [1]
           chr1 [ 1, 249250621]
           chr1 [ 1,
   [2]
                            100]
           chr1 [112, 249250621]
   [3]
           . . .
  [10]
           chr3 [ 1,
                            109]
           chr3 [121, 198022430]
  [11]
  [12]
           chr3 [ 1, 198022430]
  seqinfo: 3 sequences from an unspecified genome
```

The disjoin method represents a GRanges object as a collection of non-overlapping ranges:

```
> disjoin(g)
GRanges object with 5 ranges and 0 metadata columns:
      segnames
                   ranges strand
         <Rle> <IRanges> <Rle>
  [1]
          chr1 [101, 111]
  [2]
          chr2 [102, 102]
  [3]
          chr2 [103, 112]
          chr2 [113, 113]
  [4]
  [5]
          chr3 [110, 120]
  seqinfo: 3 sequences from an unspecified genome
```

The coverage method quantifies the degree of overlap for all the ranges in a GRanges object.

```
> coverage(g)
RleList of length 3
integer-Rle of length 249250621 with 3 runs
                100
                           11 249250510
  Lengths:
  Values :
                  0
                            1
$chr2
integer-Rle of length 243199373 with 5 runs
                 101
  Lengths:
                             1
                                      10
                                                 1 243199260
  Values :
                   0
                             1
                                       2
                                                 1
                                                           0
$chr3
integer-Rle of length 198022430 with 3 runs
                 109
  Lengths:
                            11 198022310
  Values :
```

See the *GenomicRanges* help page ?"inter-range-methods" for additional help.

Between-range methods involve operations between two *GRanges* objects; some of these are summarized in the next section.

2.4 Interval set operations for *GRanges* objects

Between-range methods calculate relationships between different GRanges objects. Of central importance are findOverlaps and related operations; these are discussed below. Additional operations treat GRanges as mathematical sets of coordinates; union(g, g2) is the union of the coordinates in g and g2. Here are examples for calculating the union, the intersect and the asymmetric difference (using setdiff).

```
> q2 < - head(qr, n=2)
> union(g, g2)
GRanges object with 3 ranges and 0 metadata columns:
                   ranges strand
      segnames
         <Rle> <IRanges> <Rle>
          chr1 [101, 111]
  [1]
  [2]
          chr2 [102, 113]
  [3]
          chr3 [110, 120]
  seqinfo: 3 sequences from an unspecified genome
> intersect(g, g2)
GRanges object with 2 ranges and 0 metadata columns:
                   ranges strand
      segnames
         <Rle> <IRanges> <Rle>
          chr1 [101, 111]
  [1]
  [2]
          chr2 [102, 112]
  seqinfo: 3 sequences from an unspecified genome
> setdiff(g, g2)
GRanges object with 2 ranges and 0 metadata columns:
      segnames
                   ranges strand
         <Rle> <IRanges> <Rle>
  [1]
          chr2 [113, 113]
          chr3 [110, 120]
  [2]
  seqinfo: 3 sequences from an unspecified genome
```

Related methods are available when the structure of the *GRanges* objects are 'parallel' to one another, i.e., element 1 of object 1 is related to element 1 of object 2, and so on. These operations all begin with a p, which is short for parallel. The methods then perform elementwise, e.g., the union of element 1 of object 1 with element 1 of object 2, etc. A requirement for these operations is that the number of elements in each *GRanges* object is the same, and that both of the objects have the same seqnames and strand assignments throughout.

```
> g3 <- g[1:2]
> ranges(g3[1]) <- IRanges(start=105, end=112)
```

```
> punion(g2, g3)
GRanges object with 2 ranges and 0 metadata columns:
                ranges strand
   segnames
      <Rle> <IRanges> <Rle>
       chr1 [101, 112]
       chr2 [102, 112]
 seqinfo: 3 sequences from an unspecified genome
> pintersect(q2, q3)
GRanges object with 2 ranges and 3 metadata columns:
   segnames
              ranges strand | score
                                                       GC
                                                               hit
      <Rle> <IRanges> <Rle> | <integer>
                                                <numeric> <logical>
                       - | 1
       chr1 [105, 111]
                                                        1
                                                                 1
       chr2 [102, 112]
                                    2 0.8888888888888
                          + |
  seqinfo: 3 sequences from an unspecified genome
> psetdiff(g2, g3)
GRanges object with 2 ranges and 0 metadata columns:
   segnames
               ranges strand
      <Rle> <IRanges> <Rle>
       chr1 [101, 104]
       chr2 [102, 101]
 seqinfo: 3 sequences from an unspecified genome
```

For more information on the GRanges classes be sure to consult the manual page.

```
> ?GRanges
```

A relatively comprehensive list of available methods is discovered with

```
> methods(class="GRanges")
```

3 GRangesList: Groups of Genomic Ranges

Some important genomic features, such as spliced transcripts that are are comprised of exons, are inherently compound structures. Such a feature makes much more sense when expressed as a compound object such as a *GRangesList*. Whenever genomic features consist of multiple ranges that are grouped by a parent feature, they can be represented as a *GRangesList* object. Consider the simple example of the two transcript GRangesList below created using the GRangesList constructor.

```
> gr1 <- GRanges(
+     seqnames = "chr2",
+     ranges = IRanges(103, 106),
+     strand = "+",
+     score = 5L, GC = 0.45)</pre>
```

```
> gr2 <- GRanges(
      seqnames = c("chr1", "chr1"),
      ranges = IRanges(c(107, 113), width = 3),
      strand = c("+", "-"),
      score = 3:4, GC = c(0.3, 0.5))
> grl <- GRangesList("txA" = gr1, "txB" = gr2)</pre>
> grl
GRangesList object of length 2:
$txA
GRanges object with 1 range and 2 metadata columns:
                   ranges strand |
                                        score
                                                     GC
         <Rle> <IRanges> <Rle> | <integer> <numeric>
          chr2 [103, 106]
  [1]
                               + |
                                            5
                                                   0.45
$txB
GRanges object with 2 ranges and 2 metadata columns:
      segnames
                   ranges strand | score GC
  [1]
          chr1 [107, 109]
                               + |
                                        3 0.3
  [2]
          chr1 [113, 115]
                               - |
                                        4 0.5
seqinfo: 2 sequences from an unspecified genome; no seqlengths
```

The show method for a *GRangesList* object displays it as a named list of *GRanges* objects, where the names of this list are considered to be the names of the grouping feature. In the example above, the groups of individual exon ranges are represented as separate *GRanges* objects which are further organized into a list structure where each element name is a transcript name. Many other combinations of grouped and labeled *GRanges* objects are possible of course, but this example is expected to be a common arrangement.

3.1 Basic GRangesList accessors

Just as with *GRanges* object, the components of the genomic coordinates within a *GRanges-List* object can be extracted using simple accessor methods. Not surprisingly, the *GRangesList* objects have many of the same accessors as *GRanges* objects. The difference is that many of these methods return a list since the input is now essentially a list of *GRanges* objects. Here are a few examples:

```
> seqnames(grl)
RleList of length 2
$txA
factor-Rle of length 1 with 1 run
    Lengths: 1
    Values : chr2
Levels(2): chr2 chr1

$txB
factor-Rle of length 2 with 1 run
    Lengths: 2
```

```
Values : chr1
Levels(2): chr2 chr1
> ranges(grl)
IRangesList of length 2
IRanges object with 1 range and 0 metadata columns:
                     end
                             width
      <integer> <integer> <integer>
  [1]
          103
                     106
$txB
IRanges object with 2 ranges and 0 metadata columns:
                     end
                             width
          start
      <integer> <integer>
  [1]
           107
                     109
                                 3
                                 3
           113
                     115
  [2]
> strand(grl)
RleList of length 2
factor-Rle of length 1 with 1 run
  Lengths: 1
  Values : +
Levels(3): + - *
$txB
factor-Rle of length 2 with 2 runs
  Lengths: 1 1
  Values : + -
Levels(3): + - *
```

The length and names methods will return the length or names of the list and the seqlengths method will return the set of sequence lengths.

```
> length(grl)
[1] 2
> names(grl)
[1] "txA" "txB"
> seqlengths(grl)
chr2 chr1
    NA NA
```

The elementNROWS method returns a list of integers corresponding to the result of calling NROW on each individual *GRanges* object contained by the *GRangesList*. This is a faster alternative to calling lapply on the *GRangesList*.

```
> elementNROWS(grl)
txA txB
```

1 2

isEmpty tests if a GRangesList object contains anything.

```
> isEmpty(grl)
[1] FALSE
```

In the context of a *GRangesList* object, the mcols method performs a similar operation to what it does on a *GRanges* object. However, this metadata now refers to information at the list level instead of the level of the individual *GRanges* objects.

Element-level metadata can be retrieved by unlisting the GRangesList, and extracting the metadata

3.2 Combining GRangesList objects

GRangesList objects can be unlisted to combine the separate GRanges objects that they contain as an expanded GRanges.

```
> ul <- unlist(grl)
> ul
GRanges object with 3 ranges and 2 metadata columns:
     seqnames
                  ranges strand |
                                     score
                                                  GC
        <Rle> <IRanges> <Rle> | <integer> <numeric>
         chr2 [103, 106]
                             + |
                                      5
                                                0.45
  txA
                              + |
         chr1 [107, 109]
                                         3
                                                 0.3
  txB
         chr1 [113, 115]
                                         4
                                                 0.5
  txB
                              - |
  seqinfo: 2 sequences from an unspecified genome; no seqlengths
```

Append lists using append or c.

A support site user had two GRangesList objects with 'parallel' elements, and wanted to combined these element-wise into a single GRangesList. One solution is to use pc() – parallel (element-wise) c(). A more general solution is to concatenate the lists and then re-group by some factor, in this case the names of the elements.

```
> grl1 <- GRangesList(</pre>
      gr1 = GRanges("chr2", IRanges(3, 6)),
      gr2 = GRanges("chr1", IRanges(c(7,13), width = 3)))
> grl2 <- GRangesList(</pre>
      gr1 = GRanges("chr2", IRanges(9, 12)),
      gr2 = GRanges("chr1", IRanges(c(25,38), width = 3)))
> pc(grl1, grl2)
GRangesList object of length 2:
$gr1
GRanges object with 2 ranges and 0 metadata columns:
      segnames
                  ranges strand
         <Rle> <IRanges> <Rle>
  [1]
          chr2
               [3, 6]
  [2]
          chr2 [9, 12]
$gr2
GRanges object with 4 ranges and 0 metadata columns:
      segnames ranges strand
  [1]
          chr1 [ 7, 9]
  [2]
          chr1 [13, 15]
  [3]
          chr1 [25, 27]
  [4]
          chr1 [38, 40]
seqinfo: 2 sequences from an unspecified genome; no seqlengths
> grl3 <- c(grl1, grl2)
> regroup(grl3, names(grl3))
GRangesList object of length 2:
GRanges object with 2 ranges and 0 metadata columns:
      segnames
                  ranges strand
         <Rle> <IRanges> <Rle>
  [1]
          chr2 [3, 6]
  [2]
          chr2 [9, 12]
$gr2
GRanges object with 4 ranges and 0 metadata columns:
      seqnames ranges strand
  [1]
          chr1 [ 7, 9]
          chr1 [13, 15]
  [2]
  [3]
          chr1 [25, 27]
  [4]
          chr1 [38, 40]
-----
seqinfo: 2 sequences from an unspecified genome; no seqlengths
```

3.3 Basic interval operations for *GRangesList* objects

For interval operations, many of the same methods exist for *GRangesList* objects that exist for *GRanges* objects.

```
> start(grl)
IntegerList of length 2
[["txA"]] 103
[["txB"]] 107 113
> end(grl)
IntegerList of length 2
[["txA"]] 106
[["txB"]] 109 115
> width(grl)
IntegerList of length 2
[["txA"]] 4
[["txB"]] 3 3
```

These operations return a data structure representing, e.g., *IntegerList*, a list where all elements are integers; it can be convenient to use mathematical and other operations on *List objects that work on each element, e.g.,

```
> sum(width(grl)) # sum of widths of each grl element
txA txB
4 6
```

Most of the intra-, inter- and between-range methods operate on *GRangesList* objects, e.g., to shift all the *GRanges* objects in a *GRangesList* object, or calculate the coverage. Both of these operations are also carried out across each *GRanges* list member.

```
> shift(grl, 20)
GRangesList object of length 2:
GRanges object with 1 range and 2 metadata columns:
     segnames
                  ranges strand | score
        <Rle> <IRanges> <Rle> | <integer> <numeric>
  [1]
         chr2 [123, 126]
                              + |
$txB
GRanges object with 2 ranges and 2 metadata columns:
     segnames
                  ranges strand | score GC
         chr1 [127, 129] + |
                                     3 0.3
  [1]
 [2]
         chr1 [133, 135]
                             - |
                                      4 0.5
seqinfo: 2 sequences from an unspecified genome; no seqlengths
> coverage(grl)
RleList of length 2
```

```
$chr2
integer-Rle of length 106 with 2 runs
  Lengths: 102  4
  Values: 0  1

$chr1
integer-Rle of length 115 with 4 runs
  Lengths: 106  3  3  3
  Values: 0  1  0  1
```

3.4 Subsetting GRangesList objects

A *GRangesList* object is behaves like a list: [returns a *GRangesList* containing a subset of the original object; [[or \$ returns the *GRanges* object at that location in the list.

```
> grl[1]
> grl[[1]]
> grl["txA"]
> grl$txB
```

In addition, subsetting a *GRangesList* also accepts a second parameter to specify which of the metadata columns you wish to select.

```
> grl[1, "score"]
GRangesList object of length 1:
GRanges object with 1 range and 1 metadata column:
      segnames
                   ranges strand |
        <Rle> <IRanges> <Rle> | <integer>
         chr2 [103, 106]
  [1]
                              + |
seqinfo: 2 sequences from an unspecified genome; no seqlengths
> grl["txB", "GC"]
GRangesList object of length 1:
GRanges object with 2 ranges and 1 metadata column:
      segnames
                   ranges strand |
        <Rle> <IRanges> <Rle> | <numeric>
  [1]
          chr1 [107, 109]
                              + |
                                         0.3
         chr1 [113, 115]
                                         0.5
  [2]
                              - |
seqinfo: 2 sequences from an unspecified genome; no seqlengths
```

The head, tail, rep, rev, and window methods all behave as you would expect them to for a list object. For example, the elements referred to by window are now list elements instead of *GRanges* elements.

```
> rep(grl[[1]], times = 3)
GRanges object with 3 ranges and 2 metadata columns:
                ranges strand | score
     segnames
                                        GC
       <Rle> <IRanges> <Rle> | <integer> <numeric>
 [1] chr2 [103, 106]
                        + | 5
     chr2 [103, 106]
                          + |
                                     5
                                           0.45
  [2]
                                    5
 [3] chr2 [103, 106]
                         + |
                                           0.45
 seqinfo: 2 sequences from an unspecified genome; no seqlengths
> rev(grl)
GRangesList object of length 2:
GRanges object with 2 ranges and 2 metadata columns:
     segnames ranges strand | score GC
       <Rle> <IRanges> <Rle> | <integer> <numeric>
  [1]
        chr1 [107, 109] + | 3 0.3
  [2]
        chr1 [113, 115]
                         - |
                                    4
                                            0.5
$txA
GRanges object with 1 range and 2 metadata columns:
                ranges strand | score GC
     segnames
 [1] chr2 [103, 106] + |
                                  5 0.45
seqinfo: 2 sequences from an unspecified genome; no seqlengths
> head(grl, n=1)
GRangesList object of length 1:
$txA
GRanges object with 1 range and 2 metadata columns:
    seqnames ranges strand | score GC
       <Rle> <IRanges> <Rle> | <integer> <numeric>
 [1]
     chr2 [103, 106] + | 5
seginfo: 2 sequences from an unspecified genome; no seglengths
> tail(grl, n=1)
GRangesList object of length 1:
GRanges object with 2 ranges and 2 metadata columns:
             ranges strand | score GC
       <Rle> <IRanges> <Rle> | <integer> <numeric>
        chr1 [107, 109] + | 3
  [1]
                                            0.3
 [2]
        chr1 [113, 115]
                         - |
                                    4
                                            0.5
seqinfo: 2 sequences from an unspecified genome; no seqlengths
> window(grl, start=1, end=1)
```

```
GRangesList object of length 1:
$txA
GRanges object with 1 range and 2 metadata columns:
                  ranges strand |
                                      score
        <Rle> <IRanges> <Rle> | <integer> <numeric>
  [1]
         chr2 [103, 106]
                              + |
                                          5
seqinfo: 2 sequences from an unspecified genome; no seqlengths
> grl[IRanges(start=2, end=2)]
GRangesList object of length 1:
$txB
GRanges object with 2 ranges and 2 metadata columns:
     seqnames ranges strand | score
        <Rle> <IRanges> <Rle> | <integer> <numeric>
         chr1 [107, 109]
                              + |
                                          3
  [1]
                                                  0.3
  [2]
         chr1 [113, 115]
                              - |
                                                  0.5
seqinfo: 2 sequences from an unspecified genome; no seqlengths
```

3.5 Looping over GRangesList objects

For *GRangesList* objects there is also a family of apply methods. These include lapply, sapply, mapply, endoapply, mendoapply, Map, and Reduce.

The different looping methods defined for *GRangesList* objects are useful for returning different kinds of results. The standard lapply and sapply behave according to convention, with the lapply method returning a list and sapply returning a more simplified output.

```
> lapply(grl, length)
$txA
[1] 1
$txB
[1] 2
> sapply(grl, length)
txA txB
1 2
```

As with *IRanges* objects, there is also a multivariate version of sapply, called mapply, defined for *GRangesList* objects. And, if you don't want the results simplified, you can call the Map method, which does the same things as mapply but without simplifying the output.

```
> grl2 <- shift(grl, 10)
> names(grl2) <- c("shiftTxA", "shiftTxB")
> mapply(c, grl, grl2)
```

```
$txA
GRanges object with 2 ranges and 2 metadata columns:
     segnames
                 ranges strand |
                                   score
        <Rle> <IRanges> <Rle> | <integer> <numeric>
        chr2 [103, 106]
                          + |
                                5
                                             0.45
 [1]
 [2]
         chr2 [113, 116]
                           + |
                                      5
                                             0.45
 seqinfo: 2 sequences from an unspecified genome; no seqlengths
$txB
GRanges object with 4 ranges and 2 metadata columns:
                 ranges strand | score
       <Rle> <IRanges> <Rle> | <integer> <numeric>
        chr1 [107, 109] + | 3
 [1]
                                      4
 [2]
         chr1 [113, 115]
                          - |
                                              0.5
                          + |
         chr1 [117, 119]
                                     3
                                              0.3
 [3]
         chr1 [123, 125] - |
 [4]
                                     4
                                              0.5
 seqinfo: 2 sequences from an unspecified genome; no seqlengths
> Map(c, grl, grl2)
$txA
GRanges object with 2 ranges and 2 metadata columns:
     segnames
                ranges strand | score
       <Rle> <IRanges> <Rle> | <integer> <numeric>
 [1]
      chr2 [103, 106]
                         + |
                                   5
                                             0.45
                                     5
                                             0.45
 [2]
     chr2 [113, 116]
                          + |
 seginfo: 2 sequences from an unspecified genome; no seglengths
GRanges object with 4 ranges and 2 metadata columns:
                 ranges strand |
                                  score
        <Rle> <IRanges> <Rle> | <integer> <numeric>
 [1]
        chr1 [107, 109]
                          + | 3
                                              0.3
         chr1 [113, 115]
 [2]
                           - |
                                     4
                                              0.5
 [3]
         chr1 [117, 119]
                           + |
                                     3
                                              0.3
 [4]
         chr1 [123, 125]
                                      4
                                              0.5
                          - 1
 seqinfo: 2 sequences from an unspecified genome; no seqlengths
```

Sometimes you will want to get back a modified version of the *GRangesList* that you originally passed in.

An endomorphism is a transformation of an object to another instance of the same class . This is achieved using the endoapply method, which will return the results as a *GRangesList* object.

```
> endoapply(grl, rev)

GRangesList object of length 2:
$txA
```

```
GRanges object with 1 range and 2 metadata columns:
                   ranges strand |
                                       score
                                                    \mathsf{GC}
      segnames
         <Rle> <IRanges> <Rle> | <integer> <numeric>
          chr2 [103, 106]
  [1]
                            + |
                                          5
$txB
GRanges object with 2 ranges and 2 metadata columns:
                  ranges strand | score GC
      segnames
          chr1 [113, 115]
                            - |
                                       4 0.5
  [1]
  [2]
          chr1 [107, 109]
                              + |
                                       3 0.3
seqinfo: 2 sequences from an unspecified genome; no seqlengths
> mendoapply(c, grl, grl2)
GRangesList object of length 2:
$txA
GRanges object with 2 ranges and 2 metadata columns:
                   ranges strand |
                                       score
      segnames
        <Rle> <IRanges> <Rle> | <integer> <numeric>
          chr2 [103, 106]
                              + |
                                         5
                                                  0.45
  [1]
                                          5
  [2]
          chr2 [113, 116]
                              + |
                                                  0.45
$txB
GRanges object with 4 ranges and 2 metadata columns:
      seqnames
                  ranges strand | score GC
  [1]
          chr1 [107, 109]
                            + |
                                       3 0.3
          chr1 [113, 115]
                              - |
                                      4 0.5
  [2]
          chr1 [117, 119]
                                      3 0.3
  [3]
                              + |
  [4]
          chr1 [123, 125]
                                       4 0.5
                              - |
seqinfo: 2 sequences from an unspecified genome; no seqlengths
```

The Reduce method will allow the *GRanges* objects to be collapsed across the whole of the *GRangesList* object.

```
> Reduce(c, grl)
GRanges object with 3 ranges and 2 metadata columns:
      segnames
                   ranges strand |
                                       score
                                                    GC
         <Rle> <IRanges> <Rle> | <integer> <numeric>
  [1]
          chr2 [103, 106]
                              + |
                                        5
                                                  0.45
  [2]
          chr1 [107, 109]
                              + |
                                          3
                                                   0.3
          chr1 [113, 115]
                                                   0.5
  [3]
                              - |
  seqinfo: 2 sequences from an unspecified genome; no seqlengths
```

Explicit element-wise operations (lapply() and friends) on *GRangesList* objects with many elements can be slow. It is therefore beneficial to explore operations that work on *List objects directly (e.g., many of the 'group generic' operators, see ?S4groupGeneric, and the

set and parallel set operators (e.g., union, punion). A useful and fast strategy is to unlist the *GRangesList* to a *GRanges* object, operate on the *GRanges* object, then relist the result, e.g.,

```
> gr <- unlist(grl)
> gr$log_score <- log(gr$score)</pre>
> grl <- relist(gr, grl)</pre>
> grl
GRangesList object of length 2:
GRanges object with 1 range and 3 metadata columns:
                  ranges strand |
                                     score GC
                                                            log_score
        <Rle> <IRanges> <Rle> | <integer> <numeric>
                                                            <numeric>
  txA
          chr2 [103, 106]
                              + |
                                          5
                                                 0.45 1.6094379124341
$txB
GRanges object with 2 ranges and 3 metadata columns:
               ranges strand | score GC
                                                   log_score
       chr1 [107, 109] + | 3 0.3 1.09861228866811
  txB
          chr1 [113, 115]
                              - |
                                      4 0.5 1.38629436111989
  txB
seqinfo: 2 sequences from an unspecified genome; no seqlengths
```

See also ?extractList.

For more information on the $\overline{GRangesList}$ classes be sure to consult the manual page and available methods

```
> ?GRangesList
> methods(class="GRangesList") # _partial_ list
```

4 Interval overlaps involving GRanges and GRanges-List objects

Interval overlapping is the process of comparing the ranges in two objects to determine if and when they overlap. As such, it is perhaps the most common operation performed on *GRanges* and *GRangesList* objects. To this end, the *GenomicRanges* package provides a family of interval overlap functions. The most general of these functions is <code>findOverlaps</code>, which takes a query and a subject as inputs and returns a *Hits* object containing the index pairings for the overlapping elements.

As suggested in the sections discussing the nature of the *GRanges* and *GRangesList* classes, the index in the above matrix of hits for a *GRanges* object is a single range while for a *GRangesList* object it is the set of ranges that define a "feature".

Another function in the overlaps family is **countOverlaps**, which tabulates the number of overlaps for each element in the query.

```
> countOverlaps(gr, grl)
txA txB txB
1 1 1
```

A third function in this family is subsetByOverlaps, which extracts the elements in the query
that overlap at least one element in the subject.

```
> subsetByOverlaps(gr,grl)
GRanges object with 3 ranges and 3 metadata columns:
      segnames
                   ranges strand |
                                      score
                                                GC
                                                             log_score
         <Rle> <IRanges> <Rle> | <integer> <numeric>
                                                             <numeric>
         chr2 [103, 106]
                              + |
                                          5
                                                 0.45 1.6094379124341
  txA
         chr1 [107, 109]
                                         3
                                                  0.3 1.09861228866811
  txB
                              + |
  txB
         chr1 [113, 115]
                              - |
                                                  0.5 1.38629436111989
  seqinfo: 2 sequences from an unspecified genome; no seqlengths
```

Finally, you can use the select argument to get the index of the first overlapping element in the subject for each element in the query.

```
> findOverlaps(gr, grl, select="first")
[1] 1 2 2
> findOverlaps(grl, gr, select="first")
[1] 1 2
```

5 Session Information

All of the output in this vignette was produced under the following conditions:

```
> sessionInfo()

R version 3.4.3 (2017-11-30)
Platform: x86_64-apple-darwin15.6.0 (64-bit)
Running under: OS X El Capitan 10.11.6

Matrix products: default
BLAS: /Library/Frameworks/R.framework/Versions/3.4/Resources/lib/libRblas.0.dylib
LAPACK: /Library/Frameworks/R.framework/Versions/3.4/Resources/lib/libRlapack.dylib

locale:
[1] C/en_US.UTF-8/en_US.UTF-8/C/en_US.UTF-8/en_US.UTF-8
attached base packages:
```

```
[1] parallel stats4
                        stats
                                  graphics grDevices utils
                                                                 datasets
[8] methods
other attached packages:
 [1] BSgenome.Scerevisiae.UCSC.sacCer2_1.4.0
 [2] KEGGgraph_1.38.0
 [3] KEGG.db_3.2.3
 [4] BSgenome.Hsapiens.UCSC.hg19_1.4.0
 [5] BSgenome_1.46.0
 [6] rtracklayer_1.38.3
 [7] edgeR_3.20.8
 [8] limma_3.34.9
 [9] DESeq2_1.18.1
[10] AnnotationHub_2.10.1
[11] TxDb.Athaliana.BioMart.plantsmart22_3.0.1
[12] TxDb.Hsapiens.UCSC.hg19.knownGene_3.2.2
[13] TxDb.Dmelanogaster.UCSC.dm3.ensGene_3.2.2
[14] GenomicFeatures_1.30.3
[15] AnnotationDbi_1.40.0
[16] GenomicAlignments_1.14.1
[17] Rsamtools_1.30.0
[18] Biostrings_2.46.0
[19] XVector_0.18.0
[20] SummarizedExperiment_1.8.1
[21] DelayedArray_0.4.1
[22] matrixStats_0.53.1
[23] Biobase_2.38.0
[24] pasillaBamSubset_0.16.0
[25] GenomicRanges_1.30.3
[26] GenomeInfoDb_1.14.0
[27] IRanges_2.12.0
[28] S4Vectors_0.16.0
[29] BiocGenerics_0.24.0
loaded via a namespace (and not attached):
 [1] bitops_1.0-6
                                   bit64_0.9-7
 [3] RColorBrewer_1.1-2
                                   progress_1.1.2
 [5] httr_1.3.1
                                   rprojroot_1.3-2
 [7] tools_3.4.3
                                   backports_1.1.2
 [9] R6_2.2.2
                                   rpart_4.1-13
[11] Hmisc_4.1-1
                                   DBI_0.7
[13] lazyeval_0.2.1
                                   colorspace_1.3-2
[15] nnet_7.3-12
                                   gridExtra_2.3
[17] prettyunits_1.0.2
                                   RMySQL_0.10.13
[19] bit_1.1-12
                                   curl_3.1
[21] compiler_3.4.3
                                   graph_1.56.0
[23] htmlTable_1.11.2
                                   scales_0.5.0
[25] checkmate_1.8.5
                                    genefilter_1.60.0
[27] stringr_1.3.0
                                   digest_0.6.15
[29] foreign_0.8-69
                                    rmarkdown_1.8
[31] base64enc_0.1-3
                                   pkgconfig_2.0.1
[33] htmltools_0.3.6
                                   htmlwidgets_1.0
[35] rlang_0.2.0
                                    rstudioapi_0.7
[37] RSQLite_2.0
                                   BiocInstaller_1.28.0
[39] shiny_1.0.5
                                   BiocParallel_1.12.0
```

VariantAnnotation_1.24.5
magrittr_1.5
Formula_1.2-2
Rcpp_0.12.15
stringi_1.1.6
zlibbioc_1.24.0
grid_3.4.3
lattice_0.20-35
annotate_1.56.1
knitr_1.20
geneplotter_1.56.0
XML_3.98-1.10
latticeExtra_0.6-28
httpuv_1.3.5
assertthat_0.2.0
mime_0.5
survival_2.41-3
memoise_1.1.0
<pre>interactiveDisplayBase_1.16.0</pre>