Bioconductor Basics: Granges and Biostrings

• Core Bioconductor structures for representing genes and genetic sequences

Motivation and Introduction

- Case study: given genomic DNA extracted from human cells, where on the genome does the nuclear protein ESRRA (estrogen related receptor alpha) bind?
- Role of estrogen receptors in breast cancer
- Data comes from analysis of ChIP-seq experiments: performed in ENCODE project import info for files in "narrowPeak" format and analyze in Bioconductor GRanges object
- Identifying nearest transcriptional start site for each binding peak assess whether regulatory activity
 of ESRRA occurs in transcriptional promoter regions

```
library(ERBS)
data(HepG2)
class(HepG2)

## [1] "GRanges"

## attr(,"package")

## [1] "GenomicRanges"
```

GenomicRanges

- ERBS library from github repo
- Load two datasets GM12878, HepG2. Estrogen receptor binding site datasets from two cell lines (cell-type dependent outcome).
- Contains: Chromosome start + end (1 row / region), strand information, score from peaks
- Access the GRanges objects as a matrix, i.e. subsetting is okay.
- \bullet sequames function to access chromosome for each row. Returns object of type Rle more efficient to save ordered by chromosome with counts. Can turn into character using as.character
- Most of analysis is focused on first 23 chromosomes
- Function to order by genomic region

Attaching package: 'BiocGenerics'

• Iranges function not specific to genomics - Granges builds on Iranges in relation to genomics

```
# install ERBS
library(devtools)

## Loading required package: usethis
install_github("genomicsclass/ERBS")

## Skipping install of 'ERBS' from a github remote, the SHA1 (9f16eb6a) has not changed since last inst
## Use `force = TRUE` to force installation
library(GenomicRanges)

## Loading required package: stats4

## Loading required package: BiocGenerics
## Loading required package: parallel
##
```

```
##
##
       clusterApply, clusterApplyLB, clusterCall, clusterEvalQ,
       clusterExport, clusterMap, parApply, parCapply, parLapply,
##
##
       parLapplyLB, parRapply, parSapply, parSapplyLB
## The following objects are masked from 'package:stats':
##
##
       IQR, mad, sd, var, xtabs
## The following objects are masked from 'package:base':
##
##
       anyDuplicated, append, as.data.frame, basename, cbind, colnames,
##
       dirname, do.call, duplicated, eval, evalq, Filter, Find, get, grep,
##
       grepl, intersect, is.unsorted, lapply, Map, mapply, match, mget,
##
       order, paste, pmax, pmax.int, pmin, pmin.int, Position, rank,
##
       rbind, Reduce, rownames, sapply, setdiff, sort, table, tapply,
##
       union, unique, unsplit, which.max, which.min
## Loading required package: S4Vectors
##
## Attaching package: 'S4Vectors'
## The following object is masked from 'package:base':
##
##
       expand.grid
## Loading required package: IRanges
## Loading required package: GenomeInfoDb
# load GM12878 and HepG2 objects from ERBS package
library(ERBS)
data(GM12878)
data(HepG2)
# inspect HepG2 GRanges object
class(HepG2)
## [1] "GRanges"
## attr(,"package")
## [1] "GenomicRanges"
HepG2
## GRanges object with 303 ranges and 7 metadata columns:
##
           seqnames
                                  ranges strand |
                                                        name
                                                                  score
                                                                               col
##
              <Rle>
                               <IRanges>
                                           <Rle> | <numeric> <integer> <logical>
##
       [1]
               chr2
                       20335378-20335787
                                               * |
                                                           NA
                                                                      0
                                                                              <NA>
##
       [2]
                           328285-329145
                                                                      0
                                                                              <NA>
              chr20
                                                           NA
##
       [3]
                                                                      0
                                                                              <NA>
               chr6 168135432-168136587
                                               * |
                                                           NA
       [4]
##
              chr19
                         1244419-1245304
                                                           NA
                                                                      0
                                                                              <NA>
##
       [5]
              chr11
                       64071828-64073069
                                               * |
                                                           NA
                                                                      0
                                                                              <NA>
##
       . . .
                . . .
                                                                               . . .
                                                          . . .
##
     [299]
                         1797182-1797852
                                                                      0
                                                                              <NA>
               chr4
                                                           NA
##
     [300]
               chr1 110198573-110199126
                                               * |
                                                           NA
                                                                      0
                                                                              <NA>
     [301]
                                                                      0
                                                                              <NA>
##
              chr17
                     17734052-17734469
                                               * |
                                                           NA
##
     [302]
               chr1
                       48306453-48306908
                                               * |
                                                           NA
                                                                      0
                                                                              <NA>
```

The following objects are masked from 'package:parallel':

```
chr12 123867207-123867554 * |
##
     [303]
                                                       NA
                                                                         <NA>
##
                         pValue
          signalValue
                                     qValue
                                                 peak
##
            <numeric> <numeric> <numeric> <integer>
##
               58.251
                         75.899 6.14371e-72
       [1]
##
       [2]
               10.808
                         69.685 5.02806e-66
                                                  321
##
       [3]
               17.103
                         54.311 7.93067e-51
                                                  930
       [4]
               12.427 43.855 1.35976e-40
##
                                                  604
       [5]
                         40.977 7.33386e-38
##
               10.850
                                                  492
##
                            ...
                  . . .
                                                  . . .
##
                         10.057 1.42334e-08
     [299]
                9.681
                                                  402
##
     [300]
                7.929
                       10.047 1.44208e-08
                                                  197
##
     [301]
                5.864
                         9.990 1.63892e-08
                                                  227
##
     [302]
                5.660
                          9.948 1.79941e-08
                                                  211
               13.211
                          9.918 1.92180e-08
                                                  163
##
     [303]
##
    seqinfo: 93 sequences (1 circular) from hg19 genome
values (HepG2)
## DataFrame with 303 rows and 7 columns
                                col signalValue
                                                   pValue
           name
                    score
                                                               qValue
                                                                           peak
##
       <numeric> <integer> <logical>
                                      <numeric> <numeric> <numeric> <integer>
## 1
             NA
                        0
                                 NA
                                         58.251
                                                   75.899 6.14371e-72
                                                                            195
## 2
             NA
                        0
                                 NA
                                         10.808
                                                   69.685 5.02806e-66
                                                                            321
## 3
             NA
                        0
                                 NA
                                         17.103
                                                   54.311 7.93067e-51
                                                                            930
                                                   43.855 1.35976e-40
## 4
             NA
                        0
                                 NA
                                         12.427
                                                                            604
## 5
             NA
                        0
                                         10.850
                                                   40.977 7.33386e-38
                                                                            492
                                 NA
             . . .
                                 . . .
                                          . . .
                                                                            . . .
## 299
                                                   10.057 1.42334e-08
             NA
                        0
                                 NA
                                          9.681
                                                                            402
## 300
             NA
                        0
                                 NA
                                          7.929
                                                   10.047 1.44208e-08
                                                                            197
## 301
                        0
             NA
                                 NA
                                          5.864
                                                   9.990 1.63892e-08
                                                                            227
## 302
             NA
                        0
                                 NA
                                          5.660
                                                    9.948 1.79941e-08
                                                                            211
                                                                            163
## 303
             NA
                        0
                                 NA
                                         13.211
                                                    9.918 1.92180e-08
# segnames extracts chromosome names
seqnames(HepG2)
                # stored as type Rle
## factor-Rle of length 303 with 292 runs
                                         1 ...
               1 1 1 1
                                                   1
    Values : chr2 chr20 chr6 chr19 chr11 ... chr4 chr1 chr17 chr1 chr12
## Levels(93): chr1 chr2 chr3 ... chrUn_gl000247 chrUn_gl000248 chrUn_gl000249
chr = seqnames(HepG2)
as.character(chr)
                    # view as character type
                "chr20" "chr6" "chr19" "chr11" "chr20" "chr19" "chr2"
##
     [1] "chr2"
                "chr6" "chr20" "chr7" "chr16" "chr9" "chr11" "chr22" "chrX"
##
    [10] "chr3"
    [19] "chr8" "chr16" "chr16" "chr19" "chr17" "chr17" "chr16" "chr16" "chr16"
##
##
    [28] "chr9"
                "chr17" "chr16" "chr12" "chr6" "chr2" "chr3" "chr11" "chr16"
##
   [37] "chr6"
                "chr2" "chr8" "chr1" "chr17" "chr20" "chr4" "chr14" "chr19"
                        "chr2" "chr2" "chr19" "chr8" "chr14" "chr22" "chr2"
    [46] "chr20" "chr9"
##
    [55] "chr14" "chr6" "chr20" "chr2" "chr19" "chr8" "chr2" "chr19" "chr12"
##
##
   [64] "chr2"
                 "chr2" "chr11" "chr12" "chr7" "chr19" "chr22" "chr17" "chr3"
   [73] "chr8"
                "chr3"
                        "chr15" "chr6" "chr9" "chr10" "chr6"
##
                                                                "chr2"
    [82] "chr11" "chr8" "chr17" "chr15" "chr21" "chr7" "chr2" "chr2"
##
    [91] "chr2" "chr16" "chr10" "chr20" "chr17" "chr13" "chr2" "chr5" "chr14"
```

```
## [100] "chr11" "chr8" "chr20" "chr3" "chr7" "chr1" "chr1" "chr3" "chr17"
## [109] "chrX" "chr19" "chr20" "chr6" "chr7" "chr16" "chr7" "chr17" "chr20"
## [118] "chr2" "chr5" "chrX" "chr7" "chr6" "chr19" "chr17" "chr16" "chr5"
## [127] "chr12" "chr9" "chr20" "chr2" "chr12" "chr3" "chr7" "chr2" "chr20"
## [136] "chr20" "chr17" "chr12" "chr19" "chr1" "chr7" "chr20" "chr14" "chr12"
## [145] "chr10" "chr6" "chr9" "chr6" "chr1" "chr18" "chr8" "chr15" "chr6"
## [154] "chr2" "chr1" "chr18" "chr16" "chr9" "chr20" "chr19" "chr17" "chr10"
## [163] "chr6" "chr2" "chrX" "chr16" "chr20" "chr16" "chr20" "chr16" "chr20"
## [172] "chr5" "chr16" "chr17" "chr17" "chr3" "chr8" "chr18" "chr18" "chr7"
## [181] "chr20" "chr16" "chr19" "chr11" "chr12" "chr2" "chr17" "chr1" "chr20"
## [190] "chr4" "chr17" "chr1" "chr6" "chr5" "chr13" "chr7" "chr20" "chr2"
## [199] "chr16" "chr6" "chr11" "chr5" "chr20" "chr1" "chr9" "chr2" "chr16"
## [208] "chr10" "chr9" "chr2" "chr2" "chr21" "chr1" "chr16" "chr18" "chr10"
## [217] "chr16" "chr3" "chr6" "chr16" "chr2" "chr6" "chr10" "chr16" "chr22"
## [226] "chr2" "chr16" "chr8" "chr20" "chr19" "chr16" "chr20" "chr2" "chr3"
## [235] "chr10" "chr14" "chr6" "chr18" "chr15" "chr9" "chr14" "chr7" "chr20"
## [244] "chr3" "chr6" "chr10" "chr4" "chr1" "chr9" "chr15" "chr6" "chr16"
## [253] "chr2" "chr3" "chr14" "chr19" "chr2" "chr5" "chr22" "chr16" "chr6"
## [262] "chr16" "chr17" "chr11" "chr8" "chr3" "chr1" "chr16" "chr21" "chr12"
## [271] "chr16" "chr1" "chr2" "chr2" "chr9" "chr2" "chr16" "chr17" "chr12"
## [280] "chr17" "chr7" "chr20" "chr7" "chr6" "chr12" "chr2" "chr1" "chr5"
## [289] "chr6" "chr2" "chr1" "chr12" "chr2" "chr6" "chr20" "chr2" "chr17"
## [298] "chr3" "chr4" "chr1" "chr17" "chr11" "chr12"
```

make a table of numbers of sequences on each chromosome table(chr)

##	chr		
##	chr1	chr2	chr3
##	18	38	15
##	chr4	chr5	chr6
##	4	8	24
##	chr7	chr8	chr9
##	14	11	12
##	chr10	chr11	chr12
##	9	9	13
##	chr13	chr14	chr15
##	2	8	5
##	chr16	chr17	chr18
##	31	21	6
##	chr19	chr20	chr21
##	16	27	3
##	chr22	chrX	chrY
##	5	4	0
##	chrM	chr1_gl000191_random	chr1_gl000192_random
##	0	0	0
##	chr4_ctg9_hap1	chr4_gl000193_random	chr4_gl000194_random
##	0	0	0
##	chr6_apd_hap1	chr6_cox_hap2	chr6_dbb_hap3
##	0	0	0
##	chr6_mann_hap4	chr6_mcf_hap5	chr6_qbl_hap6
##	0	0	0
##	chr6_ssto_hap7	chr7_gl000195_random	chr8_gl000196_random
##	0	0	0
##	chr8_gl000197_random	chr9_gl000198_random	chr9_gl000199_random

```
chr9_gl000201_random chr11_gl000202_random
##
    chr9 gl000200 random
##
         chr17_ctg5_hap1 chr17_gl000203_random chr17_gl000204_random
##
##
   chr17 gl000205 random
                          chr17 gl000206 random chr18 gl000207 random
##
##
   chr19_gl000208_random
                          chr19 gl000209 random chr21 gl000210 random
##
##
##
          chrUn_gl000211
                                  chrUn_gl000212
                                                          chrUn_gl000213
##
          chrUn_gl000214
                                  chrUn_gl000215
##
                                                          chrUn_gl000216
##
##
           chrUn_gl000217
                                  chrUn_gl000218
                                                          chrUn_gl000219
##
                                                                        0
##
           chrUn_g1000220
                                  chrUn_gl000221
                                                          chrUn_g1000222
##
                                                                        0
##
          chrUn_g1000223
                                  chrUn_gl000224
                                                          chrUn_g1000225
##
##
          chrUn_g1000226
                                  chrUn g1000227
                                                          chrUn g1000228
##
##
          chrUn_gl000229
                                  chrUn_gl000230
                                                          chrUn_gl000231
##
                        0
          chrUn_g1000232
                                  chrUn_gl000233
                                                          chrUn_gl000234
##
##
##
          chrUn_g1000235
                                  chrUn_gl000236
                                                          chrUn_gl000237
##
          chrUn_g1000238
                                  chrUn_gl000239
                                                          chrUn_g1000240
##
##
##
          chrUn_gl000241
                                  chrUn_gl000242
                                                          chrUn_g1000243
##
##
          chrUn_g1000244
                                  chrUn_gl000245
                                                          chrUn_g1000246
##
                                                          chrUn_g1000249
##
          chrUn_g1000247
                                  chrUn_gl000248
table(chr)[1:24] # restrict to autosomes, X and Y
## chr
##
                 chr3
                       chr4
                             chr5
                                    chr6
                                           chr7
                                                 chr8
                                                       chr9 chr10 chr11 chr12 chr13
                                 8
                                             14
                                                                 9
                                                                        9
             38
                   15
                                      24
                                                   11
                                                          12
   chr14 chr15 chr16 chr17 chr18 chr19
                                         chr20
                                                chr21 chr22
                                                              chrX
                                                                    chrY
                                             27
                                                    3
##
                   31
                          21
# GRanges can be subsetted and ordered
HepG2[chr=="chr20",]
   GRanges object with 27 ranges and 7 metadata columns:
##
          seqnames
                                ranges strand |
                                                      name
                                                                score
##
              <Rle>
                             <IRanges>
                                         <Rle>
                                                            <integer> <logical>
                                                 <numeric>
##
      [1]
              chr20
                        328285-329145
                                                                    0
                                                                            <NA>
                                                        NA
      [2]
                                                                    0
##
              chr20 22410891-22411863
                                                        NA
                                                                            <NA>
##
      [3]
              chr20 56039583-56040249
                                                        NA
                                                                     0
                                                                            <NA>
##
      [4]
              chr20 16455811-16456232
                                                        NA
                                                                     0
                                                                            <NA>
##
      [5]
              chr20
                      3140243-3140774
                                                        NA
                                                                     0
                                                                            <NA>
```

##

```
##
      . . .
               . . .
                                                      . . .
                                                                           . . .
             chr20
                     5591571-5592037
##
     Γ231
                                                                          <NA>
                                                       NA
                                                                  0
##
     [24]
             chr20 25519664-25520238
                                                       NA
                                                                  0
                                                                          <NA>
             chr20 19900951-19901275
     [25]
                                                                          <NA>
##
                                                       NA
                                                                  0
##
     [26]
             chr20 35156796-35157140
                                                       NA
                                                                  0
                                                                          <NA>
     [27]
             chr20 25036720-25037716
                                                       NA
                                                                          <NA>
##
                                                                  0
##
          signalValue
                         pValue
                                      qValue
##
            <numeric> <numeric>
                                   <numeric> <integer>
##
      [1]
               10.808
                          69.685 5.02806e-66
                                                    321
      [2]
##
                6.419
                          41.020 7.74961e-38
                                                    660
##
      [3]
                7.796
                          36.977 3.66693e-34
                                                    315
      [4]
                7.351
                          21.831 1.59668e-19
                                                    199
##
                7.296
##
      [5]
                          21.587 2.62536e-19
                                                    315
##
      . . .
                  . . .
                             . . .
                                                    . . .
##
     [23]
                8.766
                          11.433 7.67742e-10
                                                    249
##
     [24]
                3.300
                          11.419 7.89520e-10
                                                    206
                4.809
                          11.155 1.37954e-09
##
     [25]
                                                    140
##
     [26]
               10.154
                          10.313 8.30971e-09
                                                    163
     [27]
                4.381
                          10.087 1.33278e-08
                                                    170
##
##
     _____
##
     seqinfo: 93 sequences (1 circular) from hg19 genome
x = HepG2[order(HepG2),]
                # demonstrate usefulness of Rle type
seqnames(x)
## factor-Rle of length 303 with 23 runs
                 18
                        38
                              15
                                           8 ...
                                                     16
                                                           27
                                                                   3
##
     Lengths:
                                     4
     Values: chr1 chr2 chr3 chr4 chr5 ... chr19 chr20 chr21 chr22 chrX
## Levels(93): chr1 chr2 chr3 ... chrUn_gl000247 chrUn_gl000248 chrUn_gl000249
as.character(seqnames(x))
##
     [1] "chr1"
                 "chr1"
                          "chr1"
                                  "chr1"
                                          "chr1"
                                                   "chr1"
                                                           "chr1"
                                                                    "chr1"
                                                                            "chr1"
##
    [10] "chr1"
                 "chr1"
                          "chr1"
                                  "chr1"
                                          "chr1"
                                                   "chr1"
                                                           "chr1"
                                                                   "chr1"
                                                                            "chr1"
##
    [19] "chr2"
                  "chr2"
                          "chr2"
                                  "chr2"
                                          "chr2"
                                                   "chr2"
                                                           "chr2"
                                                                    "chr2"
                                                                            "chr2"
    [28] "chr2"
                  chr2"
                          "chr2"
                                  "chr2"
                                          "chr2"
                                                   "chr2"
                                                           "chr2"
                                                                    "chr2"
                                                                            "chr2"
##
##
    [37] "chr2"
                  "chr2"
                          "chr2"
                                  "chr2"
                                          "chr2"
                                                   "chr2"
                                                           "chr2"
                                                                    "chr2"
                                                                            "chr2"
    [46] "chr2"
                  "chr2"
                          "chr2"
                                  "chr2"
                                          "chr2"
                                                   "chr2"
                                                           "chr2"
##
                                                                    "chr2"
                                                                            "chr2"
                                                           "chr3"
##
    [55] "chr2"
                  "chr2"
                          "chr3"
                                  "chr3"
                                          "chr3"
                                                   "chr3"
                                                                    "chr3"
                                                                            "chr3"
    [64] "chr3"
                  chr3"
                          "chr3"
                                          "chr3"
                                                           "chr3"
##
                                  "chr3"
                                                   "chr3"
                                                                    "chr3"
                                                                            "chr4"
##
    [73] "chr4"
                 "chr4"
                          "chr4"
                                  "chr5"
                                          "chr5"
                                                   "chr5"
                                                           "chr5" "chr5"
                                                                            "chr5"
                 "chr5"
                                          "chr6"
##
    [82] "chr5"
                          "chr6"
                                  "chr6"
                                                   "chr6"
                                                           "chr6" "chr6"
                                                                            "chr6"
                                                   "chr6"
   [91] "chr6"
                  "chr6"
                          "chr6"
                                  "chr6"
                                          "chr6"
                                                           "chr6"
                                                                    "chr6"
##
                                                                            "chr6"
## [100] "chr6"
                                          "chr6"
                  "chr6"
                          "chr6"
                                  "chr6"
                                                   "chr6"
                                                           "chr6"
                                                                    "chr6"
                                                                            "chr7"
                          "chr7"
##
   [109] "chr7"
                 "chr7"
                                  "chr7"
                                          "chr7"
                                                   "chr7"
                                                           "chr7"
                                                                   "chr7"
                                                                            "chr7"
                          "chr7"
  [118] "chr7"
                  "chr7"
                                  "chr7"
                                          "chr8"
                                                   "chr8"
                                                           "chr8"
                                                                    "chr8"
## [127] "chr8"
                  "chr8"
                          "chr8"
                                  "chr8"
                                          "chr8"
                                                   "chr8"
                                                           "chr9"
                                                                    "chr9"
## [136] "chr9"
                 "chr9"
                          "chr9"
                                  "chr9"
                                          "chr9" "chr9"
                                                           "chr9" "chr9"
## [145] "chr10" "chr10" "chr10" "chr10" "chr10" "chr10" "chr10" "chr10" "chr10"
## [154] "chr11" "chr11" "chr11" "chr11" "chr11" "chr11" "chr11" "chr11" "chr11"
## [163] "chr12" "chr12" "chr12" "chr12" "chr12" "chr12" "chr12" "chr12" "chr12" "chr12"
## [172] "chr12" "chr12" "chr12" "chr12" "chr13" "chr13" "chr14" "chr14" "chr14"
## [181] "chr14" "chr14" "chr14" "chr14" "chr14" "chr15" "chr15" "chr15" "chr15"
## [190] "chr15" "chr16" "chr16" "chr16" "chr16" "chr16" "chr16" "chr16" "chr16"
## [199] "chr16" "chr16" "chr16" "chr16" "chr16" "chr16" "chr16" "chr16" "chr16"
```

```
## [208] "chr16" "chr17" "chr18" "chr18" "chr18" "chr18" "chr18" "chr18" "chr19" "chr20" "chr2
```

```
Assessment: Genomic Ranges
library(GenomicRanges)
paste("median of signal value column for HepG2 data: ")
## [1] "median of signal value column for HepG2 data: "
median(mcols(HepG2)$signalValue)
## [1] 7.024
paste("chromosome in region with highest signal value: ")
## [1] "chromosome in region with highest signal value: "
max_index <- which.max(mcols(HepG2)$signalValue)</pre>
chr = seqnames(HepG2)
as.character(chr)[max_index]
## [1] "chrX"
paste("Number of regions from chromosome 16: ")
## [1] "Number of regions from chromosome 16: "
HepG2[chr == "chr16",]
## GRanges object with 31 ranges and 7 metadata columns:
##
          segnames
                               ranges strand
                                                     name
                                                               score
                                                                            col
##
             <Rle>
                            <IRanges> <Rle> | <numeric> <integer> <logical>
             chr16 70191209-70192150
##
      [1]
                                                       NA
                                                                   0
                                                                           <NA>
      [2]
             chr16 1701039-1702137
                                                                           <NA>
##
                                                       NA
                                                                   0
      [3]
##
             chr16 25189109-25190026
                                                                   0
                                                                           <NA>
                                                       NA
      [4]
             chr16 85325101-85325686
##
                                                       NA
                                                                   0
                                                                           <NA>
      [5]
             chr16 29986461-29986872
##
                                                       NA
                                                                   0
                                                                           <NA>
##
      . . .
                                                                           . . .
##
     [27]
             chr16 57481218-57481854
                                                                   0
                                                                           <NA>
                                                       NA
##
     [28]
             chr16 85322504-85322950
                                                                   0
                                                                           <NA>
                                                       NA
##
     [29]
             chr16 19134897-19135280
                                                       NA
                                                                   0
                                                                           <NA>
##
     [30]
             chr16
                      2586101-2586737
                                                       NA
                                                                   0
                                                                           <NA>
##
     [31]
             chr16 29975932-29976255
                                                                   0
                                                                           <NA>
##
          signalValue
                          pValue
                                       qValue
                                                   peak
##
            <numeric> <numeric>
                                   <numeric> <integer>
##
      [1]
                8.371
                          37.774 8.19277e-35
                                                    688
```

36.264 1.65696e-33

##

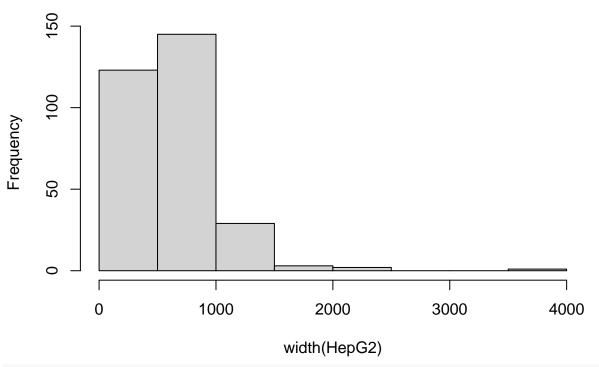
[2]

16.157

```
[3]
                                                      606
##
                 5.979
                           31.808 3.44356e-29
      [4]
##
                 7.664
                           31.429 7.88321e-29
                                                      223
      [5]
                14.795
##
                           29.018 1.73008e-26
                                                      198
##
##
     [27]
                 5.126
                           10.761 3.20978e-09
                                                      196
##
     [28]
                 4.331
                           10.725 3.43494e-09
                                                      223
##
     [29]
                 5.380
                           10.562 4.92563e-09
                                                      203
                                                      472
##
     [30]
                 6.521
                           10.514 5.42123e-09
##
     [31]
                 6.897
                           10.436 6.37196e-09
                                                      145
##
     seqinfo: 93 sequences (1 circular) from hg19 genome
```

hist(width(HepG2))

Histogram of width(HepG2)



```
median_width <- median(width(HepG2))
paste("Median width of all chromosomes: ", median_width)</pre>
```

[1] "Median width of all chromosomes: 560"

Bioconductor Infrastructure for genomics, microarray and NGS

- IRanges package representing ranges of integers. Base pair arrangements we want to manipulate in genomics
- Vignette about classes and functions in IRanges package
- Simple functions have good performance
- Summary of most important functions
- IRanges start, end, width (i.e., 5, 10, 6bP long)

- Start, end, and width functions
- Can specify > 1 range at a time to make IRanges objects of length n
- Intra-range methods:
- Shift Intra range methods for IRanges doesn't depend on other ranges contained in IRanges object. I.e., shift IRange to the left by 2.
- Narrow relative to start, start at nth base pair
- Flank get flanking sequence 3 base pairs from start or end (start = False). Also bi-directional (both=True)
- Inter-range methods:
- range will give beginning of the IRanges to the end, including gaps in between
- reduce gives us base pairs covered by the original ranges (do not get gaps). Can ask for gaps.
- **disjoint** set of ranges which has the same coverage as original IRanges object but non-overlapping. Contain union of all endpoints of the original range.

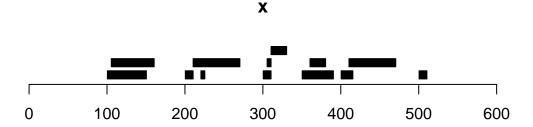
Assessment: IRanges

```
library(IRanges)
ir <- IRanges(101, 200)
paste("*2 zooms in, giving range with half the width. New starting point: ", start(ir*2))
## [1] "*2 zooms in, giving range with half the width. New starting point: 126"
n_ir <- narrow(ir, start=20)</pre>
paste("narrow function with start of 20. New starting point: ", start(n_ir))
## [1] "narrow function with start of 20. New starting point: 120"
paste("+25 operation gives width of resulting range: ", width(ir+25))
## [1] "+25 operation gives width of resulting range: 150"
m_ir <- IRanges(start=c(1, 11, 21),end=c(3, 15, 27))</pre>
paste("sum of widths of multiple IRanges objects:", sum(width(m_ir)))
## [1] "sum of widths of multiple IRanges objects: 15"
library(ph525x)
## Loading required package: png
## Loading required package: grid
## Loading required package: Biobase
## Welcome to Bioconductor
##
      Vignettes contain introductory material; view with
##
      'browseVignettes()'. To cite Bioconductor, see
##
      'citation("Biobase")', and for packages 'citation("pkgname")'.
##
## Loading required package: Homo.sapiens
## Loading required package: AnnotationDbi
```

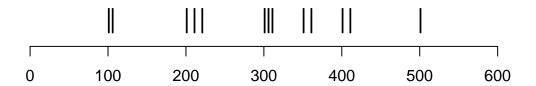
```
## Loading required package: OrganismDbi
## Loading required package: GenomicFeatures
## Loading required package: GO.db
##
## Loading required package: org.Hs.eg.db
## Loading required package: TxDb.Hsapiens.UCSC.hg19.knownGene
plotRanges(x)
                                  X
100
                200
                                 300
                                                 400
                                                                 500
paste("Total width not covered by ranges in x:", sum(width(gaps(x))))
## [1] "Total width not covered by ranges in x: 130"
paste("Number of disjoint ranges within ranges in x:", length(disjoin(x)))
## [1] "Number of disjoint ranges within ranges in x: 17"
par(mfrow=c(2, 1))
```

plotRanges(x, xlim=c(0, 600))

plotRanges(resize(x, 1), xlim=c(0, 600))







Genomic ranges: GRanges

- Extension of IRanges
- Contain a sequence name IRange of chromosome Z.
- Can contain chromosome information and sequence length
- Sequence names as Rle
- IRanges and strange as Rle also
- Can shift similar to IRanges will go off end of chromosome if exceeds length
- Wrap in trim function to make sure that the end at chromosome end does not exceed
- \bullet Metadata accessed with mcols
- Can add cols by mcols\$
- Additional package called GRangesList groups GRanges together by wrapping in function call
- Example of GRangesList grouping exons by gene or by transcript
- Application of package find overlaps between GRanges objects
- findOverlaps function query and subject (see in help() function)
- output of $\mathit{findOverlaps}$ is a hits object with length representing # overlaps
- Same way to get the overlaps is %over% function which returns logical vector
- \bullet Rle object defined by IRanges but similar object in base R=Run length encoding
- If vector repeats certain values, can save memory by number and number of times repeated
- str function gives us the compact representation
- Peering into *Rle* object can use *Views* object to see *IRanges* from start to end. Only a virtual class saves *Rle* and number of views / windows into it
- Can also use for Fasta files or other objects

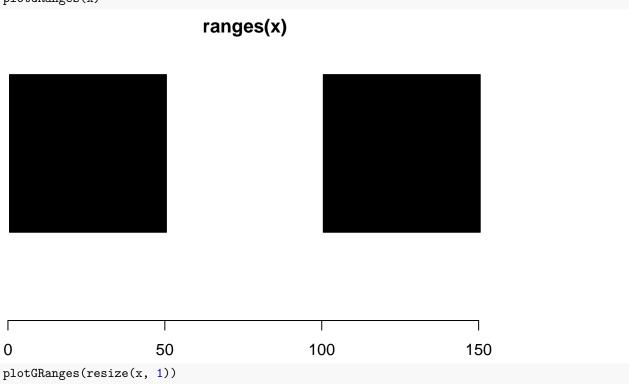
Assessment: GRanges

- GRanges object extends concept of interval ranges
- Ranges can be defined by:
 - chromosome we are referring to (segnames in Bioconductor)
 - strand of DNA we are referring to (+ or -)

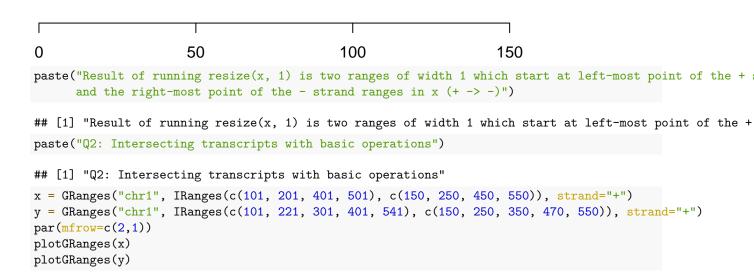
• These two pieces of information are necessary for specification of a range of DNA

```
library(GenomicRanges)
library(IRanges)
library(ph525x)
x = GRanges("chr1", IRanges(c(1,101),c(50,150)), strand=c("+","-"))
paste("Get the internal IRanges from a GRanges object: ")

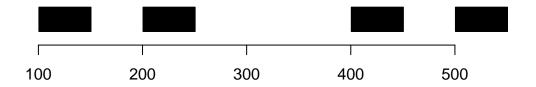
## [1] "Get the internal IRanges from a GRanges object: "
plotGRanges = function(x) plotRanges(ranges(x))
plotGRanges(x)
```



ranges(x)



ranges(x)



ranges(x)

```
100
               200
                               300
                                              400
                                                              500
multiple_granges_list <- GRangesList(x,y)</pre>
single_granges_list <- GRangesList(c(x, y))</pre>
findOverlaps(x, y)
## Hits object with 4 hits and 0 metadata columns:
##
         queryHits subjectHits
##
         <integer>
                      <integer>
##
     [1]
##
     [2]
                 2
                              2
     [3]
                 3
                              4
##
                 4
                              5
##
     [4]
##
     queryLength: 4 / subjectLength: 5
##
paste("width of overlap between x and y: ", sum(width(union(x, y))) - sum(width(intersect(x, y))))
## [1] "width of overlap between x and y: 130"
z = GRanges("chr1", range(ranges(x)), strand="-")
```