PeakSegDisk usage examples

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Welcome to PeakSegDisk, an R package for optimal peak detection in very large count data sequences.

1 Related work

The PeakSeg R packages contain algorithms for inferring optimal segmentation models subject to the constraint that up changes must be followed by down changes, and vice versa. This ensures that the model can be interpreted in terms of peaks (after up changes) and background (after down changes).

- PeakSegDP the historically first PeakSeg package, https://CRAN.R-project.org/package=PeakSegDP provides a heuristic quadratic time algorithm for computing models from 1 to S segments for a single sample. This was the original algorithm described in our ICML'15 paper, http://jmlr.org/proceedings/papers/v37/hocking15.html, but it is neither fast nor optimal, so in practice we recommend to use our newer packages below instead.
- PeakSegOptimal https://CRAN.R-project.org/package=PeakSegOptimal provides log-linear time algorithms for computing optimal models with multiple peaks for a single sample. The algorithms are faster and more accurate than PeakSegDP, https://arxiv.org/abs/1703.03352
- PeakSegDisk https://github.com/tdhock/PeakSegDisk provides an on-disk implementation of optimal log-linear algorithms for computing multiple peaks in a single sample. Computes same models as PeakSegOptimal but works for much larger data sets because disk is used for storage instead of memory. https://arxiv.org/abs/1810.00117
- PeakSegJoint https://CRAN.R-project.org/package=PeakSegJoint provides a fast heuristic algorithm for computing models with a single common peak in 0, ..., S samples. https://arxiv.org/abs/1506.01286
- PeakSegPipeline https://github.com/tdhock/PeakSegPipeline provides a pipeline for genome-wide peak calling using the other PeakSeg packages.

The remainder of this vignette is dedicated to an explanation of how to use PeakSegDisk.

2 Simulate a noisy integer vector with changes

The first example we will treat is detecting peaks in a vector of integer data, with possibly the same values at adjacent positions. This is an inefficient representation for large genomic data, but it is the typical output from simulation functions like rpois:

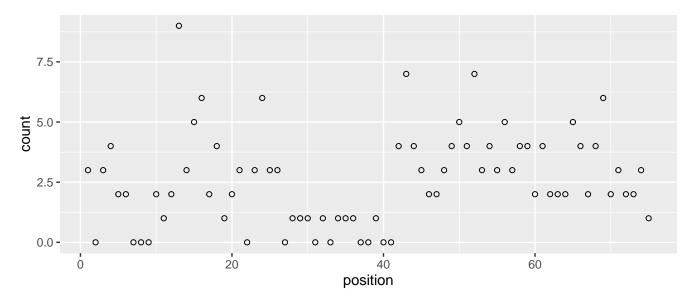
```
sim.seg <- function(seg.mean, size.mean=15) {
    seg.size <- rpois(1, size.mean)
    rpois(seg.size, seg.mean)
}
set.seed(1)
seg.mean.vec <- c(1.5, 3.5, 0.5, 4.5, 2.5)
z.list <- lapply(seg.mean.vec, sim.seg)
(z.rep.vec <- unlist(z.list))

#> [1] 3 0 3 4 2 2 0 0 0 2 1 2 9 3 5 6 2 4 1 2 3 0 3 6 3 3 0 1 1 1 0 1 0 1 1 1 0 0
#> [39] 1 0 0 4 7 4 3 2 2 3 4 5 4 7 3 4 3 5 3 4 4 2 4 2 2 2 5 4 2 4 6 2 3 2 2 3 1
```

From the output above it is clear that these simulated data are integers, with some identical values at adjacent positions.

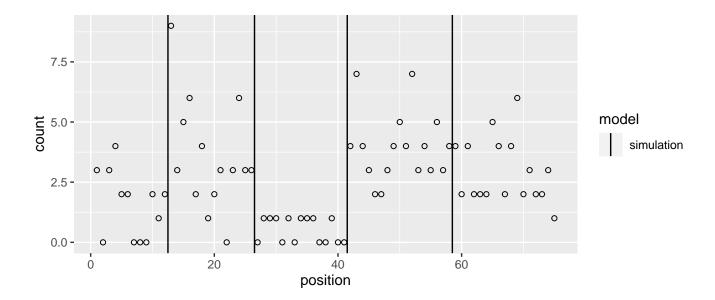
Below we put these data into a data table in order to plot them along with the model using ggplot2:

```
count.df <- data.frame(
   chrom="chrUnknown",
   chromStart=0:(length(z.rep.vec)-1),
   chromEnd=1:length(z.rep.vec),
   count=z.rep.vec)
if(require(ggplot2)){
gg.count <- ggplot()+
   xlab("position")+
   geom_point(aes(
        chromEnd, count),
        shape=1,
        data=count.df)
gg.count
}
#> Loading required package: ggplot2
```



The true changepoints in the simulation are shown below.

```
n.segs <- length(seg.mean.vec)</pre>
seg.size.vec <- sapply(z.list, length)</pre>
seg.end.vec <- cumsum(seg.size.vec)</pre>
change.vec <- seg.end.vec[-n.segs]+0.5
change.df <- data.frame(</pre>
  changepoint=change.vec)
if(require(ggplot2)){
gg.change <- gg.count+
  geom_vline(aes(
    xintercept=changepoint, color=model),
    data=data.frame(change.df, model="simulation"))+
  scale_color_manual(
    values=c(
      simulation="black",
      fitted="green"))
gg.change
```



3 Segment a vector of integers

Let $z_1, \ldots, z_n \in \mathbb{Z}_+$ be the sequence of n non-negative count data in z.rep.vec, and let $w_1 = \cdots = w_n = 1$ be weights which are all 1. The peak detection algorithm computes the solution to the following optimization problem:

where $\ell(m, z) = m - z \log m$ is the Poisson loss. The optimization variables are m_i for the segment mean, s_i for hidden state, and c_i for type of changepoint. The penalty term is proportional to the number of changepoint variables c_i which are equal to 1 (which is the same as the number of peaks in the resulting model).

To run the peak detection algorithm a numeric penalty parameter $\lambda \geq 0$ must be specified by the user. The smallest value is 0 which yields max peaks, and the largest value is Inf which yields no peaks. The code below runs the peak detection algorithm on this count data vector, using the penalty parameter $\lambda = 10.5$:

```
fit <- list()
(fit$vec <- PeakSegDisk::PeakSegFPOP_vec(z.rep.vec, 10.5))

#> $segments
#> chrom chromStart chromEnd status mean
#> <char> <int> <char> <num>
```

```
#> 1: chrUnknown
                                  75 background 2.166670
                         69
#> 2: chrUnknown
                         41
                                  69 peak 3.714290
#> 3: chrUnknown
                         26
                                  41 background 0.533333
#> 4: chrUnknown
                                          peak 3.571430
                         12
                                  26
#> 5: chrUnknown
                                  12 background 1.583330
                         0
#>
#> $loss
     penalty segments peaks bases bedGraph.lines mean.pen.cost total.loss
        <num>
                 <int> <int> <int>
                                            <int>
                                                           <num>
#>
                                                                      <num>
        10.5
                     5
                         2
                                75
                                               60
                                                      0.01507491 -19.86938
      equality.constraints mean.intervals max.intervals megabytes seconds
#>
                     <int>
                                    <num>
                                                 <int>
                                                              <num>
#> 1:
                         0
                                      4.6
                                                       8 0.01372147
                                                                          0
#>
#> $data
           chrom chromStart chromEnd count
#>
          <char>
                       <int>
                                <int> <int>
                                    1
   1: chrUnknown
                           0
#>
#>
   2: chrUnknown
                           1
                                    2
                                          0
   3: chrUnknown
                           2
                                    3
#>
#> 4: chrUnknown
                           3
                                    4
#> 5: chrUnknown
                          4
                                    6
#> 6: chrUnknown
                          6
                                    9
                                          0
#> 7: chrUnknown
                          9
                                   10
#> 8: chrUnknown
                          10
                                   11
                                          1
#> 9: chrUnknown
                          11
                                   12
                                          2
#> 10: chrUnknown
                         12
                                   13
#> 11: chrUnknown
                                          3
                         13
                                   14
#> 12: chrUnknown
                          14
                                   15
#> 13: chrUnknown
                          15
                                          6
                                   16
#> 14: chrUnknown
                          16
                                   17
#> 15: chrUnknown
                          17
                                   18
                                          4
#> 16: chrUnknown
                          18
                                   19
                                          1
#> 17: chrUnknown
                          19
                                   20
                                          2
#> 18: chrUnknown
                          20
                                   21
#> 19: chrUnknown
                          21
                                   22
                                          0
#> 20: chrUnknown
                          22
                                   23
                                          3
#> 21: chrUnknown
                          23
                                          6
                                   24
#> 22: chrUnknown
                          24
                                   26
                                          3
#> 23: chrUnknown
                          26
                                   27
                                          \cap
#> 24: chrUnknown
                          27
                                          1
                                   30
#> 25: chrUnknown
                          30
                                   31
                                          0
#> 26: chrUnknown
                                   32
                                          1
                          31
#> 27: chrUnknown
                          32
                                   33
                                          0
#> 28: chrUnknown
                          33
                                   36
                                          1
#> 29: chrUnknown
                          36
                                   38
```

```
30: chrUnknown
                            38
                                      39
#> 31: chrUnknown
                            39
                                      41
                                              0
#> 32: chrUnknown
                            41
                                      42
                                              4
#> 33: chrUnknown
                                              7
                            42
                                      43
#> 34: chrUnknown
                            43
                                      44
                                              4
#> 35: chrUnknown
                            44
                                      45
                                              3
#> 36: chrUnknown
                                              2
                            45
                                      47
#> 37: chrUnknown
                            47
                                      48
                                              3
#> 38: chrUnknown
                                              4
                            48
                                      49
#> 39: chrUnknown
                            49
                                      50
                                              5
                                              4
#> 40: chrUnknown
                            50
                                      51
#> 41: chrUnknown
                            51
                                      52
                                              7
#> 42: chrUnknown
                            52
                                      53
                                              3
#> 43: chrUnknown
                            53
                                      54
                                              4
#> 44: chrUnknown
                            54
                                      55
                                              3
                                              5
#> 45: chrUnknown
                            55
                                      56
#> 46: chrUnknown
                            56
                                              3
                                      57
#> 47: chrUnknown
                                              4
                            57
                                      59
#> 48: chrUnknown
                            59
                                      60
                                              2
#> 49: chrUnknown
                            60
                                      61
                                              4
#> 50: chrUnknown
                                              2
                            61
                                      64
#> 51: chrUnknown
                                              5
                            64
                                      65
#> 52: chrUnknown
                            65
                                      66
                                              4
#> 53: chrUnknown
                                              2
                            66
                                      67
#> 54: chrUnknown
                            67
                                              4
                                      68
#> 55: chrUnknown
                                              6
                            68
                                      69
                                              2
#> 56: chrUnknown
                            69
                                      70
                                              3
#> 57: chrUnknown
                            70
                                      71
                                              2
#> 58: chrUnknown
                            71
                                      73
#> 59: chrUnknown
                            73
                                      74
                                              3
#> 60: chrUnknown
                            74
                                      75
                                              1
#>
             chrom chromStart chromEnd count
#>
#> attr(,"class")
#> [1] "PeakSegFPOP_df"
                           "PeakSegFPOP_dir" "list"
```

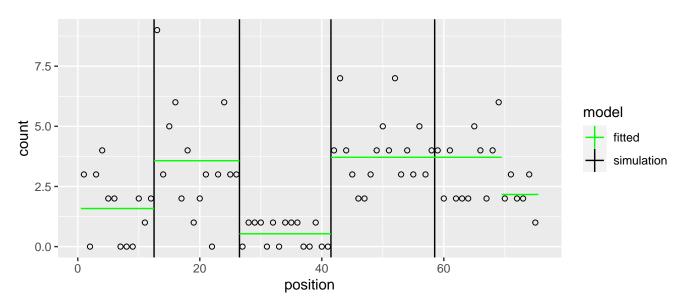
The model output list above includes **segments**, a data table with one row for each segment mean, and **loss**, a data table with one row that reports the model meta-data. Of interest are:

- penalty, the user-provided penalty value,
- segments, the number of segments,
- peaks, the number of peaks (even-numbered segments),
- bases, the number of data points in repetitive form (not run-length encoding),
- bedGraph.lines, the number of data points in run-length encoding form,

- mean.pen.cost, the optimal mean loss plus penalty*peaks,
- total.loss, the optimal total Poisson loss over all data points,
- equality.constraints, the number of adjacent segment means that are equal in the optimal solution. Note that when this number is greater than 0, then there are some active equality constraints, and the optimal model is therefore not feasible for the strict inequality constraints, which implies that the optimum of the problem with strict inequality constraints is undefined, i.e. for any sub-optimal solution that satisfies the strict inequality constraints, we can find a lower cost solution that satisfies the strict inequality constraints (but is still sub-optimal), by getting closer to the solution with active equality constraints.
- megabytes, the storage space on disk used by the solver,
- seconds, the amount of time used by the solver,
- mean.intervals, max.intervals, statistics over all intervals (candidate changepoints) computed by the functional pruning algorithm, useful for analyzing computational complexity, which is linear in the number of intervals.

Note in particular that PeakSegFPOP_vec internally uses rle to construct a run-length encoding, which is passed to the solver to save time/storage. In this case the repetitive integer data vector contains 75 elements but the coverage.bedGraph data file contains only 60 lines. In real genomic data sets the difference is typically much larger.

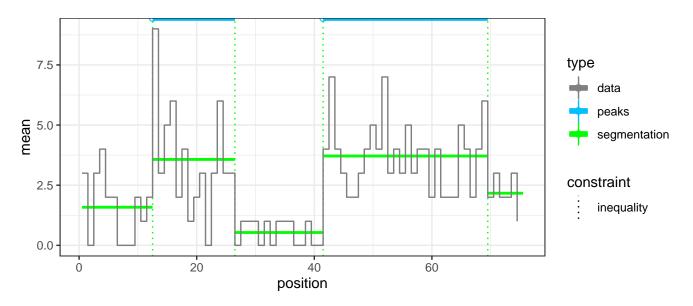
```
gg.change+
  geom_segment(aes(
    chromStart+0.5, mean, xend=chromEnd+0.5, yend=mean, color=model),
  data=data.frame(fit$vec$segments, model="fitted"))
```



It is clear from the plot above that the first three changepoints are estimated exactly and the last one is a bit over-estimated.

Also note that a default plot method is defined for these objects:

plot(fit\$vec)



4 Segment a data frame

Another interface that can be used on a data frame with n rows and exactly 4 columns (chrom, chrom-Start, chromEnd, count) is PeakSegFPOP_df. For each row $i \in \{1, ..., n\}$, let $z_i \in \mathbb{Z}_+$ be the non-negative count data (count column), and let $w_i > 0$ be the weight (equal to the number of bases, chromEnd-chromStart). The optimization problem we solve is the same as before. Note that this function does not perform run-length encoding for you:

```
(fit$df <- PeakSegDisk::PeakSegFPOP_df(count.df, 10.5))</pre>
#> $segments
#>
           chrom chromStart chromEnd
                                            status
                                                       mean
#>
          <char>
                       <int>
                                 <int>
                                            <char>
                                                      <num>
#> 1: chrUnknown
                          69
                                    75 background 2.166670
#> 2: chrUnknown
                          41
                                    69
                                              peak 3.714290
  3: chrUnknown
                                    41 background 0.533333
                          26
#> 4: chrUnknown
                          12
                                    26
                                              peak 3.571430
  5: chrUnknown
                                    12 background 1.583330
                           0
#>
#> $loss
#>
      penalty segments peaks bases bedGraph.lines mean.pen.cost total.loss
#>
        <num>
                  <int> <int> <int>
                                               <int>
                                                              <num>
                                                         0.01507491 -19.86938
         10.5
                      5
                            2
#> 1:
                                  75
                                                  75
      equality.constraints mean.intervals max.intervals megabytes seconds
#>
#>
                      <int>
                                      <num>
                                                     <int>
                                                                 <num>
                                                                          <num>
#> 1:
                          0
                                   4.633333
                                                         8 0.01725006
                                                                           0.01
#>
#> $data
```

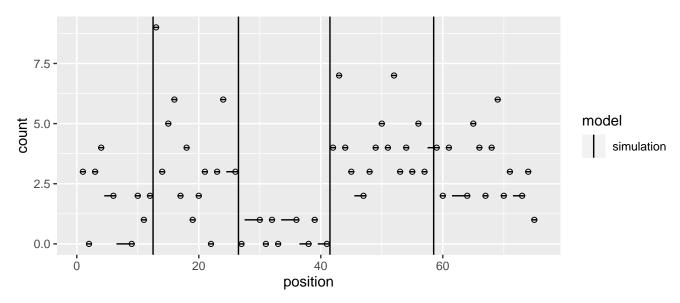
| 4~ | | a b ··· | ahmamatarrt | ahmamEm 1 | 001111 |
|----|-----|--------------------|-------------|-----------|-------------|
| #> | | | chromStart | | |
| #> | 4 | <char></char> | <int></int> | | <int></int> |
| #> | 1: | chrUnknown | 0 | 1 | 3 |
| #> | 2: | | 1 | 2 | 0 |
| #> | 3: | chrUnknown | 2 | 3 | 3 |
| #> | 4: | ${\tt chrUnknown}$ | 3 | 4 | 4 |
| #> | 5: | chrUnknown | 4 | 5 | 2 |
| #> | 6: | chrUnknown | 5 | 6 | 2 |
| #> | | chrUnknown | 6 | 7 | 0 |
| #> | | chrUnknown | 7 | 8 | 0 |
| #> | | chrUnknown | 8 | 9 | |
| | | | | | 0 |
| #> | | chrUnknown | 9 | 10 | 2 |
| #> | | chrUnknown | 10 | 11 | 1 |
| #> | | chrUnknown | 11 | 12 | 2 |
| #> | 13: | ${\tt chrUnknown}$ | 12 | 13 | 9 |
| #> | 14: | chrUnknown | 13 | 14 | 3 |
| #> | 15: | chrUnknown | 14 | 15 | 5 |
| #> | 16: | chrUnknown | 15 | 16 | 6 |
| #> | | chrUnknown | 16 | 17 | 2 |
| #> | | chrUnknown | 17 | 18 | 4 |
| | | | | | |
| #> | | chrUnknown | 18 | 19 | 1 |
| #> | | chrUnknown | 19 | 20 | 2 |
| #> | | chrUnknown | 20 | 21 | 3 |
| #> | 22: | chrUnknown | 21 | 22 | 0 |
| #> | 23: | ${\tt chrUnknown}$ | 22 | 23 | 3 |
| #> | 24: | chrUnknown | 23 | 24 | 6 |
| #> | 25: | chrUnknown | 24 | 25 | 3 |
| #> | 26: | chrUnknown | 25 | 26 | 3 |
| #> | | chrUnknown | 26 | 27 | 0 |
| | | chrUnknown | 27 | 28 | 1 |
| | | | | | |
| | | chrUnknown | 28 | 29 | 1 |
| | | chrUnknown | 29 | 30 | 1 |
| | | chrUnknown | 30 | 31 | 0 |
| #> | 32: | chrUnknown | 31 | 32 | 1 |
| #> | 33: | chrUnknown | 32 | 33 | 0 |
| #> | 34: | chrUnknown | 33 | 34 | 1 |
| #> | 35: | chrUnknown | 34 | 35 | 1 |
| | | chrUnknown | 35 | 36 | 1 |
| | | chrUnknown | 36 | 37 | 0 |
| | | | | | |
| | | chrUnknown | 37 | 38 | 0 |
| | | chrUnknown | 38 | 39 | 1 |
| | | chrUnknown | 39 | 40 | 0 |
| #> | 41: | chrUnknown | 40 | 41 | 0 |
| #> | 42: | ${\tt chrUnknown}$ | 41 | 42 | 4 |
| #> | 43: | chrUnknown | 42 | 43 | 7 |
| #> | 44: | chrUnknown | 43 | 44 | 4 |
| | | | | | |

```
#> 45: chrUnknown
                                      45
                            44
                                              3
                            45
                                              2
#> 46: chrUnknown
                                      46
                                              2
#> 47: chrUnknown
                            46
                                      47
#> 48: chrUnknown
                            47
                                              3
                                      48
#> 49: chrUnknown
                                              4
                            48
                                      49
#> 50: chrUnknown
                            49
                                      50
                                              5
#> 51: chrUnknown
                            50
                                      51
                                              4
                                              7
#> 52: chrUnknown
                            51
                                      52
#> 53: chrUnknown
                            52
                                      53
                                              3
#> 54: chrUnknown
                                              4
                            53
                                      54
#> 55: chrUnknown
                                              3
                            54
                                      55
#> 56: chrUnknown
                            55
                                      56
                                              5
#> 57: chrUnknown
                            56
                                      57
                                              3
#> 58: chrUnknown
                                              4
                            57
                                      58
#> 59: chrUnknown
                            58
                                      59
                                              4
#> 60: chrUnknown
                                              2
                            59
                                      60
#> 61: chrUnknown
                            60
                                      61
                                              4
#> 62: chrUnknown
                                              2
                                      62
                            61
#> 63: chrUnknown
                            62
                                      63
                                              2
#> 64: chrUnknown
                                              2
                            63
                                      64
#> 65: chrUnknown
                                              5
                            64
                                      65
#> 66: chrUnknown
                                              4
                            65
                                      66
#> 67: chrUnknown
                            66
                                      67
                                              2
#> 68: chrUnknown
                                              4
                            67
                                      68
#> 69: chrUnknown
                            68
                                      69
                                              6
#> 70: chrUnknown
                            69
                                      70
                                              2
#> 71: chrUnknown
                                              3
                            70
                                      71
#> 72: chrUnknown
                                      72
                                              2
                            71
#> 73: chrUnknown
                            72
                                      73
                                              2
#> 74: chrUnknown
                            73
                                      74
                                              3
#> 75: chrUnknown
                            74
                                      75
                                              1
#>
             chrom chromStart chromEnd count
#>
#> attr(,"class")
#> [1] "PeakSegFPOP_df" "PeakSegFPOP_dir" "list"
```

Note how bedGraph.lines is now the same size as bases, 75. The time/storage complexity is log-linear in the number of bedGraph.lines, so it is more efficient to use the run-length encoding. This can be easily done in R:

```
z.rle.vec <- rle(z.rep.vec)
chromEnd <- cumsum(z.rle.vec$lengths)
rle.df <- data.frame(
   chrom="chrUnknown",
   chromStart=c(OL, chromEnd[-length(chromEnd)]),
   chromEnd,</pre>
```

```
count=z.rle.vec$values)
if(require(ggplot2)){
gg.rle <- ggplot()+
  geom_segment(aes(
    chromStart+0.5, count, xend=chromEnd+0.5, yend=count),
    data=rle.df)+
  geom_point(aes(
    chromEnd, count),
    shape=1,
    data=rle.df)+
  geom_vline(aes(
    xintercept=changepoint, color=model),
    data=data.frame(change.df, model="simulation"))+
  scale_color_manual(
    values=c(
      simulation="black",
      fitted="green"))+
  xlab("position")
gg.rle
```

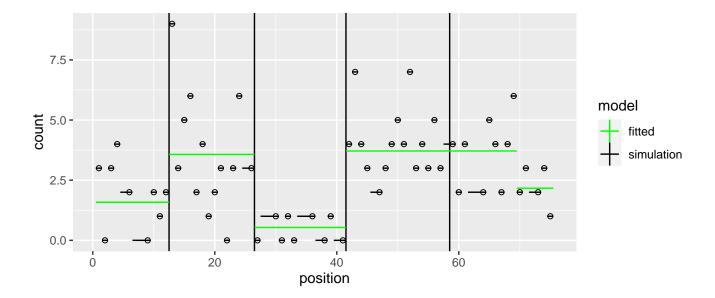


The plot above shows the run-length encoded data, with a geom_point for the last position in each run, and a geom_segment extending left to the first position. These data can be segmented as above:

```
(fit$rle <- PeakSegDisk::PeakSegFPOP_df(rle.df, 10.5))</pre>
#> $segments
#>
            chrom chromStart chromEnd
                                             status
                                                        mean
#>
           <char>
                        <int>
                                  <int>
                                             <char>
                                                        <num>
#> 1: chrUnknown
                           69
                                     75 background 2.166670
#> 2: chrUnknown
                           41
                                               peak 3.714290
```

```
#> 3: chrUnknown
                                    41 background 0.533333
                          26
#> 4: chrUnknown
                          12
                                    26
                                            peak 3.571430
#> 5: chrUnknown
                           0
                                    12 background 1.583330
#>
#> $loss
#>
      penalty segments peaks bases bedGraph.lines mean.pen.cost total.loss
                  <int> <int> <int>
#>
        <num>
                                              <int>
                                                             <num>
         10.5
#> 1:
                      5
                            2.
                                 75
                                                  60
                                                        0.01507491 - 19.86938
      equality.constraints mean.intervals max.intervals megabytes seconds
#>
#>
                      <int>
                                      <num>
                                                     <int>
                                                                 <num>
#> 1:
                          0
                                        4.6
                                                         8 0.01372147
                                                                          0.02
#>
#> $data
            chrom chromStart chromEnd count
#>
           <char>
                        <int>
                                  <int> <int>
    1: chrUnknown
                            0
                                      1
                                      2
#>
    2: chrUnknown
                            1
                                            0
    3: chrUnknown
                            2
                                      3
#>
#>
   4: chrUnknown
                            3
                                      4
                                            4
   5: chrUnknown
                            4
                                      6
                                            2
#>
   6: chrUnknown
                            6
                                      9
                                            \cap
#>
                            9
                                            2
   7: chrUnknown
                                     10
#>
#> 8: chrUnknown
                           10
                                     11
                                            1
                                            2
#> 9: chrUnknown
                           11
                                     12
#> 10: chrUnknown
                           12
                                     13
#> 11: chrUnknown
                           13
                                     14
                                            3
#> 12: chrUnknown
                                            5
                           14
                                     15
#> 13: chrUnknown
                           15
                                     16
                                            6
#> 14: chrUnknown
                           16
                                     17
#> 15: chrUnknown
                           17
                                            4
                                     18
#> 16: chrUnknown
                           18
                                     19
                                            1
#> 17: chrUnknown
                                     20
                                            2
                           19
#> 18: chrUnknown
                                            3
                           20
                                     21
#> 19: chrUnknown
                           21
                                     22
                                            0
#> 20: chrUnknown
                                            3
                           22
                                     23
#> 21: chrUnknown
                           23
                                     24
                                            6
#> 22: chrUnknown
                                            3
                           24
                                     26
#> 23: chrUnknown
                                            0
                           26
                                     27
#> 24: chrUnknown
                           27
                                     30
                                            1
#> 25: chrUnknown
                           30
                                     31
                                            ()
#> 26: chrUnknown
                                     32
                                            1
                           31
#> 27: chrUnknown
                           32
                                     33
                                            0
#> 28: chrUnknown
                                            1
                           33
                                     36
#> 29: chrUnknown
                           36
                                     38
                                            0
#> 30: chrUnknown
                           38
                                     39
                                             1
#> 31: chrUnknown
                           39
                                     41
                                            0
```

```
#> 32: chrUnknown
                                     42
                           41
#> 33: chrUnknown
                           42
                                     43
                                             7
#> 34: chrUnknown
                           43
                                     44
                                             4
#> 35: chrUnknown
                           44
                                     45
                                             3
#> 36: chrUnknown
                           45
                                     47
                                             2
                                             3
#> 37: chrUnknown
                           47
                                     48
#> 38: chrUnknown
                                             4
                           48
                                     49
#> 39: chrUnknown
                                             5
                           49
                                     50
#> 40: chrUnknown
                                             4
                           50
                                     51
#> 41: chrUnknown
                                             7
                           51
                                     52
#> 42: chrUnknown
                           52
                                     53
                                             3
#> 43: chrUnknown
                           53
                                     54
                                             4
#> 44: chrUnknown
                           54
                                     55
                                             3
#> 45: chrUnknown
                                             5
                           55
                                     56
#> 46: chrUnknown
                           56
                                     57
                                             3
#> 47: chrUnknown
                                             4
                           57
                                     59
#> 48: chrUnknown
                                             2
                           59
                                     60
#> 49: chrUnknown
                           60
                                     61
                                             4
#> 50: chrUnknown
                           61
                                     64
                                             2
#> 51: chrUnknown
                           64
                                     65
                                             5
#> 52: chrUnknown
                                     66
                                             4
                           65
                                             2
#> 53: chrUnknown
                           66
                                     67
#> 54: chrUnknown
                           67
                                     68
                                             4
#> 55: chrUnknown
                                     69
                                             6
                           68
#> 56: chrUnknown
                           69
                                     70
                                             2
#> 57: chrUnknown
                           70
                                     71
                                             3
#> 58: chrUnknown
                                             2
                           71
                                     73
#> 59: chrUnknown
                           73
                                     74
                                             3
#> 60: chrUnknown
                           74
                                     75
#>
            chrom chromStart chromEnd count
#>
#> attr(,"class")
#> [1] "PeakSegFPOP_df"
                          "PeakSegFPOP_dir" "list"
if(require(ggplot2)){
gg.rle+
  geom_segment(aes(
    chromStart+0.5, mean, xend=chromEnd+0.5, yend=mean, color=model),
    data=data.frame(fit$rle$segments, model="fitted"))
```



5 Write the file yourself

The interfaces discussed in the previous sections are perhaps the most intuitive for useRs, but they are also the least efficient, so they are not recommended for large data.

In this section we introduce the most efficient way of using PeakSegDisk, which involves:

- creating a "problem" directory for each segmentation problem (sample and genome subset),
- saving the data to coverage.bedGraph in that directory,
- and then running PeakSegFPOP_dir.

The reason why this method is recommended for large data is because PeakSegFPOP_dir saves its results to the "problem" directory. So if a certain result has already been computed, these result files are used as a cache, and are read instead of doing computations, which saves a lot of time. The file system is used as the interface in order to support very large data sets with very little memory usage.

To use PeakSegFPOP_dir the data should be saved to a chrXX-start-end/coverage.bedGraph file, where the problem directory "chrXX-start-end" should be named using a genome postion string:

- chrXX is the chromosome (which is irrelevant to the algorithm),
- start is the 0-based first position of the region to segment (the smallest possible value is 0),
- end is the 1-based end position (the smallest possible value is 1).

```
data.dir <- file.path(
   tempfile(),
   with(rle.df, sprintf(
     "%s-%d-%d", chrom[1], min(chromStart), max(chromEnd))))
dir.create(data.dir, showWarnings=FALSE, recursive=TRUE)
coverage.bedGraph <- file.path(data.dir, "coverage.bedGraph")</pre>
```

```
write.table(
  rle.df, coverage.bedGraph,
  sep="\t", row.names=FALSE, col.names=FALSE)
```

The next step is to run the main solver,

```
(fit$dir <- PeakSegDisk::PeakSegFPOP_dir(data.dir, 10.5))
#> $segments
#>
           chrom chromStart chromEnd
                                           status
                                                      mean
#>
          <char>
                       <int>
                                <int>
                                           <char>
                                                     <num>
#> 1: chrUnknown
                          69
                                   75 background 2.166670
#> 2: chrUnknown
                          41
                                   69
                                             peak 3.714290
#> 3: chrUnknown
                                   41 background 0.533333
                          26
#> 4: chrUnknown
                                             peak 3.571430
                          12
                                   26
#> 5: chrUnknown
                           0
                                   12 background 1.583330
#>
#> $loss
      penalty segments peaks bases bedGraph.lines mean.pen.cost total.loss
#>
        <num>
                  <int> <int> <int>
                                              <int>
                                                             <num>
                                                                         <num>
         10.5
#> 1:
                      5
                            2
                                                 60
                                                       0.01507491 -19.86938
#>
      equality.constraints mean.intervals max.intervals megabytes seconds
#>
                      <int>
                                      <num>
                                                    <int>
                                                                <num>
#> 1:
                          0
                                        4.6
                                                         8 0.01372147
                                                                          0.02
#>
#> attr(,"class")
#> [1] "PeakSegFPOP_dir" "list"
```

The underlying C++ code creates penalty-specific files such as

chrXX-start-end/coverage.bedGraph_penalty=0.1_loss.tsv which are used to store/cache the results. If the files already exist (and are consistent) then PeakSegFPOP_dir just reads them; otherwise it runs the dynamic programming C++ code in order to create those files, which are then read into R.

6 Computing the model with a given number of peaks

The **sequentialSearch_dir** function can be used to compute the optimal model with a certain number of peaks:

```
if(interactive() && requireNamespace("future"))future::plan("multisession")
(fit$search <- PeakSegDisk::sequentialSearch_dir(data.dir, 2L, verbose=1))

#> Loading required namespace: future.apply

#> Next = 0, Inf

#> Next = 2.20991803112367

#> Next = 7.11950550040458
```

```
#> $segments
#>
           chrom chromStart chromEnd
                                            status
                                                        mean
          <char>
                       <int>
                                 <int>
#>
                                            <char>
                                                       <num>
#> 1: chrUnknown
                           69
                                    75 background 2.166670
#> 2: chrUnknown
                           41
                                              peak 3.714290
                                    41 background 0.533333
#> 3: chrUnknown
                           26
#> 4: chrUnknown
                           12
                                    26
                                              peak 3.571430
#> 5: chrUnknown
                            0
                                    12 background 1.583330
#>
#> $loss
       penalty segments peaks bases bedGraph.lines mean.pen.cost total.loss
#>
         <num>
                   <int> <int> <int>
                                                <int>
                                                                <num>
                                                                            <num>
  1: 7.119506
                       5
                              2
                                   75
                                                    60
                                                         -0.07507161
                                                                      -19.86938
      equality.constraints mean.intervals max.intervals megabytes seconds
#>
#>
                      <int>
                                       <num>
                                                      <int>
                                                                          <num>
#> 1:
                           ()
                                   4.558333
                                                          9 0.0136261
                                                                          0.02
#>
      iteration under
                        over
#>
          <num> <int> <int>
               3
                     0
#> 1:
#>
#> $others
#>
       penalty segments peaks bases bedGraph.lines mean.pen.cost total.loss
                   <int> <int> <int>
#>
         <num>
                                                <int>
                                                                <num>
#> 1: 0.000000
                      53
                             26
                                   75
                                                    60
                                                         -0.63772877 -47.829658
#> 2:
                       1
                              0
                                   75
                                                    60
                                                          0.12837615
           Inf
                                                                        9.628211
#> 3: 2.209918
                      13
                              6
                                   75
                                                    60
                                                         -0.26439085 -33.088822
#> 4: 7.119506
                       5
                              2
                                   75
                                                         -0.07507161 -19.869382
                                                    60
      equality.constraints mean.intervals max.intervals
#>
                                                              megabytes seconds
#>
                      <int>
                                       <num>
                                                      <int>
                                                                   <num>
                                                                            <num>
#> 1:
                           5
                                   2.550000
                                                          4 0.009029388
                                                                             0.02
#> 2:
                           0
                                                          0 0.000000000
                                   0.000000
                                                                            0.00
                           0
#> 3:
                                   4.708333
                                                          9 0.013969421
                                                                             0.02
#> 4:
                           0
                                   4.558333
                                                          9 0.013626099
                                                                            0.02
#>
      iteration under
#>
          <num> <int> <int>
#> 1:
               1
                    NA
                           NA
#> 2:
               1
                    NΑ
                           NΑ
               2
#> 3:
                     0
                           26
#> 4:
               3
                     0
                            6
#>
#> attr(,"class")
#> [1] "PeakSegFPOP_dir" "list"
```

The algorithm must evaluate several penalty values to compute the optimal model with a certain number of peaks. The others component of the model list above shows that

• the search starts with penalty values 0 and Inf, which result in models with 26 and 0 peaks,

respectively.

- the next penalty evaluated is 2.21, which results in 6 peaks.
- the final penalty evaluated is 7.12, which results in 2 peaks.

At each step (except the first) the new penalties are computed based on the loss values found in the previous step. If present with a registered parallel future plan, the future.apply package is used to run the first step (penalties $0, \infty$) in parallel.

Note how the number of peaks and total.loss of this model is the same as the other models computed above,

```
lossDF <- function(L)data.frame(L$loss)[, names(fit$dir$loss)]</pre>
do.call(rbind, lapply(fit, lossDF))
#>
             penalty segments peaks bases bedGraph.lines mean.pen.cost total.loss
           10.500000
                                   2
                                                                            -19.86938
#> vec
                             5
                                         75
                                                         60
                                                                0.01507491
                             5
                                   2
#> df
          10.500000
                                         75
                                                         75
                                                               0.01507491
                                                                            -19.86938
#> rle
          10.500000
                             5
                                   2
                                         75
                                                               0.01507491
                                                                            -19.86938
                                                         60
#> dir
          10.500000
                             5
                                   2
                                         75
                                                         60
                                                               0.01507491
                                                                            -19.86938
#> search 7.119506
                             5
                                   2
                                         75
                                                         60
                                                              -0.07507161
                                                                            -19.86938
#>
           equality.constraints mean.intervals max.intervals
                                                                 megabytes seconds
                                                              8 0.01372147
#> vec
                               0
                                        4.600000
                                                                                0.00
                               0
                                        4.633333
                                                              8 0.01725006
                                                                                0.01
#> df
#> rle
                                        4.600000
                               0
                                                              8 0.01372147
                                                                                0.02
#> dir
                               0
                                        4.600000
                                                              8 0.01372147
                                                                                0.02
#> search
                                        4.558333
                                                              9 0.01362610
                                                                                0.02
```

Finally we demonstrate how the filesystem caching is especially useful for the sequential search. In the code below we ask the sequential search algorithm to compute the optimal model with four peaks:

```
four.peaks <- PeakSegDisk::sequentialSearch_dir(data.dir, 4L)</pre>
four.peaks$others[, .(iteration, penalty, peaks)]
#>
      iteration
                  penalty peaks
                     <num> <int>
#>
           <num>
#> 1:
               1 0.000000
                               26
#> 2:
               1
                       Inf
                                0
#> 3:
               2 2.209918
                                6
                                2
#> 4:
               3 7.119506
#> 5:
               4 3.304860
                                3
#> 6:
               5 2.830674
                                5
#> 7:
               6 3.107790
                                4
```

Looking at the output above, we see that the first three iterations of the sequential search require computing models with 26, 0, 6, 2 peaks. Since all of these have been previously computed (and saved to disk), the dynamic programming algorithm does not need to be re-run, and instead the model results are simply read from the files. After that the dynamic programming is run for the subsequent iterations 4-6. In this particular example the savings in computation time is not extraordinary, but in real genomic data, this can result in substantial speed-ups.