LTMG - GCR track (LTMG->biclustering analysis)

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Analysis

We will have a data namely "Yan" for the example of LTMG - GCR (biclustering) pipeline. Basically, we may need the following steps for this analysis

(i) a standard data loading function

```
data0 <- log(as.matrix(read.delim("Yan_expression_RPKM.txt", row.names = 1)))</pre>
```

(ii) running LTMG -> a standard output of LTMG parameters

Select genes

Genes with non-zero expression in more than 5 samples in data0

```
selected.genes <- which(rowSums(data0 > 0) > 5)
print(head(selected.genes, 30))
```

##	RPS11	ELM02	CREB3L1	PNMA1	TMEM216	L0C653712
##	2	3	4	5	7	8
##	C10orf90	ZHX3	ERCC5	APBB2	PDCL3	AEN
##	9	10	11	14	17	18
##	FRG2	DECR1	SALL1	RPS18	SLC10A7	BRIX1
##	19	20	21	24	25	27
##	LMAN1	CHD8	SUM01	GP1BA	UQCR11	DDB1
##	28	29	30	31	32	33
##	MYO9B	CRNKL1	XAB2	RTN1	UTY	CENPQ
##	34	35	37	38	42	43

Run LTMG for the selected genes

LTMG -> output list(N: number of peaks; a $3 \times N$ matrix: A, U, S; If Zcut (If 0 expression is more than 5, Zcut); Iteration number, upper limit 1000)

```
library(LTMGSCA)
for (gene in head(selected.genes, 3)) {
  for (k in 1:5) {
    print(SeparateKRpkmNew(x = data0[gene, ], n = 100, q = 0, k = k, err = 1e-10))
  }
}
```

```
##
                                 mean
## Late_blastocyst.3_Cell.5 1 7.74797 0.5936434
                                    p
## Late_blastocyst.2_Cell.7 0.5189909 7.632955 0.5688547
  X4.cell embryo.1 Cell.4 0.4810091 7.872066 0.5948461
##
                                    р
## X2.cell embryo.1 Cell.2 0.1210234 7.083230 0.1913294
## Late blastocyst.3 Cell.5 0.7937210 7.768004 0.5549263
## X4.cell_embryo.3_Cell.2  0.0852556  8.505077  0.1026225
                                      p
## Late_blastocyst.2_Cell.4 0.006933978 7.788798 0.36486339
## Morulae.1_Cell.8
                            0.448993364 7.267528 0.35678557
## Morulae.2_Cell.3
                            0.480661380 8.095959 0.44529688
## X4.cell_embryo.3_Cell.1
                           0.063411278 8.507573 0.09591338
##
                                      p
                                            mean
## Oocyte.1
                            0.005328433 7.727174 0.3884293
## Late_blastocyst.2_Cell.7 0.356379577 7.199494 0.3289284
## Late blastocyst.3 Cell.5 0.309216553 7.755904 0.3974516
## X4.cell_embryo.1_Cell.4   0.322192465   8.343583   0.3577630
## X8.cell_embryo.1_Cell.1 0.006882971 7.925339 0.4211202
##
                                 mean
                                           sd
## X8.cell_embryo.3_Cell.1 1 1.558705 2.07965
##
                                                     sd
                                   р
## X8.cell embryo.2 Cell.4 0.3138683 -2.385534 2.460532
## X8.cell_embryo.1_Cell.1 0.6861317 2.645441 1.280540
                                           mean
                                    p
## Late_blastocyst.2_Cell.1 0.2726636 -2.943190 1.0155455
## X8.cell_embryo.3_Cell.1 0.4874510
                                       1.851239 0.8095599
## X4.cell_embryo.1_Cell.2 0.2398854
                                       4.123094 0.1710736
##
                                   p
                                          mean
                                                      sd
## X8.cell_embryo.2_Cell.6 0.2722165 -2.584755 0.8242646
## X8.cell_embryo.2_Cell.7 0.2930781 1.687644 0.7978886
## X8.cell_embryo.2_Cell.3 0.1945976 2.090904 0.7665579
## X2.cell_embryo.3_Cell.1 0.2401079 4.123058 0.1711315
                                           mean
                                    р
                           0.27343650 -2.346844 0.7156853
## Morulae.1 Cell.2
## X8.cell embryo.2 Cell.4 0.22096460
                                      1.561465 0.7070268
## X8.cell_embryo.3_Cell.1 0.16654932 1.775654 0.7409112
## X8.cell_embryo.1_Cell.1 0.09679837 2.609410 0.5179847
## X4.cell_embryo.2_Cell.4 0.24225121
                                      4.122422 0.1716394
                                    mean
  Late blastocyst.2 Cell.3 1 -0.9535114 1.562651
##
                                           mean
                                   p
## Zygote.1
                           0.8524664 -1.4311960 1.4985531
## X8.cell_embryo.1_Cell.2 0.1475336 0.9262817 0.9096018
##
                                     р
## X4.cell_embryo.2_Cell.3 0.70557942 -1.2865546 0.4785381776
## Late_blastocyst.2_Cell.3 0.27219997
                                        0.7239671 0.5938110246
## X4.cell_embryo.3_Cell.3 0.02222061
                                        2.7078834 0.0005667612
                                            mean
## Zygote.2
                           0.72496505 -1.2936882 0.4602185469
## X2.cell_embryo.2_Cell.1 0.11346035 0.4516975 0.2993225110
## X8.cell_embryo.1_Cell.4 0.13935362 1.0987104 0.4882202184
## X4.cell embryo.1 Cell.2 0.02222099 2.7078834 0.0005667612
```

Here we have the BIC functions:

```
BIC_f_zcut <- function(y, rrr, Zcut) {</pre>
  n <- length(y)</pre>
  nparams <- nrow(rrr) * 3</pre>
  w <- rrr[, 1]
  u <- rrr[, 2]
  sig <- rrr[, 3]
  cc <- c()
  y0 \leftarrow y[which(y \ge Zcut)]
  y1 <- y[which(y < Zcut)]
  y1 <- y1 * 0 + Zcut
  for (i in 1:nrow(rrr)) {
    c0 <- dnorm(y0, u[i], sig[i]) * w[i]</pre>
    c1 <- (1 - pnorm(y1, u[i], sig[i])) * w[i]
    c <- c(c0, c1)
    cc <- rbind(cc, c)</pre>
  }
  d <- apply(cc, 2, sum)</pre>
  e <- sum(log(d))
  f \leftarrow e * 2 - nparams * log(n)
  return (f)
}
BIC_f_zcut2 <- function(y, rrr, Zcut) {</pre>
  n <- length(y)</pre>
  nparams <- nrow(rrr) * 3</pre>
  w <- rrr[, 1]
  u <- rrr[, 2]
  sig <- rrr[, 3]
  y0 \leftarrow y[which(y \ge Zcut)]
  cc <- c()
  for (i in 1:nrow(rrr)) {
    c <- dnorm(y0, u[i], sig[i]) * w[i]</pre>
    cc <- rbind(cc, c)</pre>
  }
  d <- apply(cc, 2, sum)</pre>
  e <- sum(log(d))
  f <- e * 2 - nparams * log(n)
  return (f)
}
```

We can now get f value using BIC_f_zcut2().

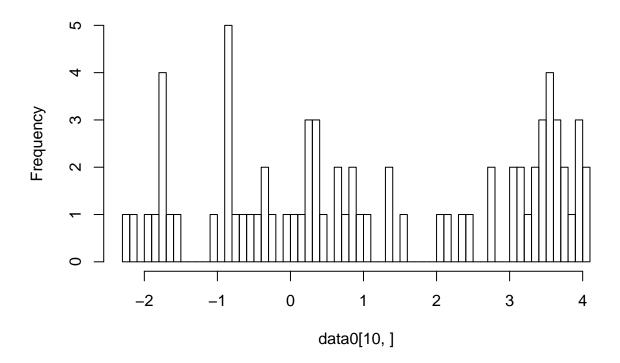
```
for (k in 1:5) {
  rrr <- SeparateKRpkmNew(x = data0[selected.genes[1], ], n = 100, q = 0, k = k, err = 1e-10)
  print(BIC_f_zcut2(y = data0[selected.genes[1], ], rrr, 0))
}
## [1] -175.0426
## [1] -188.4811
## [1] -195.0944
## [1] -208.8539
## [1] -223.8346
We are only print while k > 1.
GetBestK <- function(x, n, q, err = 1e-10){</pre>
  best.bic <- -Inf</pre>
  best.k <- 0
  best.result <- c(0, 0, 0)
  for (k in 1:7) {
    rrr <- SeparateKRpkmNew(x = x, n = n, q = q, k = k, err = err)
    bic <- BIC_f_zcut2(y = x, rrr, q)
    if(is.nan(bic)) {
     bic <- -Inf
    }
    if (bic >= best.bic) {
      best.bic <- bic</pre>
      best.k <- k
     best.result <- rrr
    } else {
      return(list(k = best.k, bic = best.bic, result = best.result))
  }
  return(list(k = 0, bic = 0, result = c(0, 0, 0)))
for (gene in head(selected.genes, 30)) {
  best <- GetBestK(x = data0[gene, ], n = 100, q = 0, err = 1e-10)
  if (best[1] > 1) {
    print(gene)
  }
}
## [1] 10
## [1] 11
## [1] 14
## [1] 17
## [1] 29
## [1] 35
```

This is the 10th one:

```
best <- GetBestK(x = data0[10, ], n = 100, q = 0, err = 1e-10)
print(best)</pre>
```

hist(data0[10,], breaks = 60)

Histogram of data0[10,]



(iii) LTMG -> discretization

We have the following functions ready for this step:

- calculate_prob_sep_Zcut,
- 2. discretization_method_1_LLR_mean, and
- $3. \ {\tt Build_R_matrix}.$

```
calculate_prob_sep_Zcut <- function(data1, Zcut, a, u, sig) {</pre>
  cc <- matrix(0, length(a), length(data1))</pre>
  colnames(cc) <- names(data1)</pre>
  for (i in 1:length(a)) {
    c <- a[i] / sig[i] * exp(-(data1 - u[i]) ^ 2 / (2 * sig[i] ^ 2))
    cc[i,] <- c
  cut_p <- rep(0, length(a))</pre>
  for (i in 1:length(a)) {
    cut_p[i] <- a[i] * pnorm(Zcut, u[i], sig[i])</pre>
  for (i in 1:ncol(cc)) {
    if (data1[i] < Zcut) {</pre>
      cc[, i] <- cut_p
    }
  }
  cc[which(is.na(cc) == 1)] <- 0
  return(cc)
}
```

```
discretization_method_1_LLR_mean <- function(y, aaa, ccc, LLR_cut = 2) {</pre>
  K <- 1 / LLR cut + 1
  if (nrow(aaa) == 1) {
    print("Only one class")
    return(y)
  } else {
    discretized_y <- rep(0, length(y))</pre>
    for (i in 1:ncol(ccc)) {
      11 \leftarrow which(ccc[, i] = max(ccc[, i]))[1]
      if ((max(ccc[, i])/sum(ccc[, i])) > (1/K)) {
        discretized_y[i] <- 11</pre>
      }
    }
    blocks <- c()
    st_c <- 1
    end_c <- 1
    st_c_v \leftarrow y[order(y)[1]]
    end_c_v \leftarrow y[order(y)[1]]
    label_c <- discretized_y[order(y)[1]]</pre>
    for (i in 2:length(order(y))) {
      if (discretized_y[order(y)[i]] == discretized_y[order(y)[i - 1]]) {
        end_c <- i
        end_c_v <- y[order(y)[i]]</pre>
        if (i == length(order(y))) {
           end c \leftarrow i
           end_c_v <- y[order(y)[i]]</pre>
           blocks <- rbind(blocks, c(st_c, end_c, st_c_v, end_c_v, label_c))</pre>
        }
      } else {
        blocks <- rbind(blocks, c(st_c, end_c, st_c_v, end_c_v,</pre>
           label_c))
        label_c <- discretized_y[order(y)[i]]</pre>
        st_c <- i
```

```
end_c <- i
        st_c_v <- y[order(y)[i]]
        end_c_v <- y[order(y)[i]]</pre>
        if (i == length(order(y))) {
          end_c <- i
          end_c_v <- y[order(y)[i]]</pre>
          blocks <- rbind(blocks, c(st_c, end_c, st_c_v, end_c_v, label_c))</pre>
      }
    }
    if (nrow(blocks) > 1) {
      for (i in 1:nrow(blocks)) {
        if (blocks[i, 5] != 0) {
          tg_i <- blocks[i, 5]
          if (!((blocks[i, 3] <= aaa[tg_i, 2]) & (blocks[i, 4] >= aaa[tg_i, 2]))) {
          blocks[i, 5] <- 0
        }
      }
      for (i in 1:nrow(blocks)) {
        discretized_y[order(y)[blocks[i, 1]:blocks[i, 2]]] <- blocks[i, 5]</pre>
    }
    return(discretized_y)
  }
}
```

```
Build_R_matrix <- function(cc, Zcut0, U, Gname) {
   tg_s <- intersect(which(U > Zcut0), unique(cc))
   dd <- c()
   nc <- c()
   if (length(tg_s) > 0) {
      for (i in 1:length(tg_s)) {
        nc <- c(nc, paste(Gname, tg_s[i], sep = "__"))
        ccc <- (cc == tg_s[i]) * 1
      dd <- rbind(dd, ccc)
    }
}
rownames(dd) <- nc
return(dd)
}</pre>
```

best\$result is a $K \times 3$ matrix with 1st, 2nd and 3rd columns are the A, U, S of the gene x is the normalized expression level

```
i <- 4
x <- data0[i, ]
Zcut0 <- 0
best <- GetBestK(x = x, n = 1000, q = Zcut0, err = 1e-10)

pp <- calculate_prob_sep_Zcut(x, Zcut0, best$result[, 1], best$result[, 2], best$result[, 3])
cc <- discretization_method_1_LLR_mean(x, best$result, pp, LLR_cut = 0.1)

## [1] "Only one class"</pre>
```

dd <- Build_R_matrix(cc, Zcut0, best\$result[, 2], rownames(data0)[i]) print(x)</pre>

```
##
                                                 Oocvte.2
                     Oocyte.1
##
                    0.000000
                                                0.3534698
##
                     Oocyte.3
                                                 Zygote.1
##
                    0.6657760
                                                0.5905606
##
                     Zygote.2
                                                 Zygote.3
##
                    0.3611648
                                               -0.3133418
##
     X2.cell_embryo.1_Cell.1
                                 X2.cell_embryo.1_Cell.2
##
                    0.4643627
                                                0.2342813
##
     X2.cell_embryo.2_Cell.1
                                 X2.cell_embryo.2_Cell.2
##
                    0.6339278
                                                0.3708737
##
     X2.cell_embryo.3_Cell.1
                                 X2.cell_embryo.3_Cell.2
##
                   -0.3523984
                                               -0.7052198
##
     X4.cell_embryo.1_Cell.1
                                 X4.cell_embryo.1_Cell.2
##
                          -Inf
                                                1.5526559
##
     X4.cell_embryo.1_Cell.3
                                 X4.cell_embryo.1_Cell.4
##
                          -Inf
                                                     -Inf
##
     X4.cell_embryo.2_Cell.1
                                 X4.cell_embryo.2_Cell.2
##
                   -1.3586792
                                               -0.2943711
##
     X4.cell_embryo.2_Cell.3
                                 X4.cell_embryo.2_Cell.4
##
                    0.4491630
                                                1.0217312
##
     X4.cell_embryo.3_Cell.1
                                 X4.cell_embryo.3_Cell.2
##
                    1.1177611
                                                0.9250522
##
     X4.cell embryo.3 Cell.3
                                 X4.cell_embryo.3_Cell.4
##
                    1.4548872
                                                1.6122340
##
     X8.cell_embryo.1_Cell.1
                                 X8.cell_embryo.1_Cell.2
##
                   -0.7635696
                                                1.0328285
##
     X8.cell_embryo.1_Cell.3
                                 X8.cell_embryo.1_Cell.4
##
                    0.6559644
                                                0.9266370
     X8.cell_embryo.2_Cell.1
                                 X8.cell_embryo.2_Cell.2
##
##
                         -Inf
                                                     -Inf
     X8.cell_embryo.2_Cell.3
##
                                 X8.cell_embryo.2_Cell.4
##
                         -Inf
                                                     -Inf
##
     X8.cell_embryo.2_Cell.5
                                 X8.cell_embryo.2_Cell.6
##
                         -Inf
                                                     -Inf
##
     X8.cell_embryo.2_Cell.7
                                 X8.cell_embryo.2_Cell.8
##
                         -Inf
                                                     -Inf
##
     X8.cell_embryo.3_Cell.1
                                 X8.cell_embryo.3_Cell.2
##
                         -Inf
                                                     -Inf
##
     X8.cell_embryo.3_Cell.3
                                 X8.cell_embryo.3_Cell.4
##
                         -Inf
                                                     -Inf
##
     X8.cell_embryo.3_Cell.5
                                 X8.cell_embryo.3_Cell.6
##
                         -Inf
                                                     -Tnf
##
     X8.cell_embryo.3_Cell.7
                                 X8.cell_embryo.3_Cell.8
##
                         -Inf
                                                1.7516317
##
            Morulae.1_Cell.1
                                        Morulae.1_Cell.2
##
                         -Inf
                                                     -Inf
##
            Morulae.1_Cell.3
                                        Morulae.1_Cell.4
##
                          -Inf
                                               -0.2930297
##
            Morulae.1_Cell.5
                                        Morulae.1_Cell.6
```

```
##
                         -Inf
                                                    -Inf
##
            Morulae.1_Cell.7
                                       Morulae.1_Cell.8
##
                        -Inf
##
            Morulae.2_Cell.1
                                       Morulae.2_Cell.2
##
                         -Inf
                                                    -Inf
            Morulae.2 Cell.3
                                       Morulae.2 Cell.4
##
##
                         -Inf
                                                    -Inf
##
            Morulae.2 Cell.5
                                       Morulae.2 Cell.6
##
                         -Inf
                                              -0.9702191
##
            Morulae.2_Cell.7
                                       Morulae.2_Cell.8
##
                  -0.9597203
                                                    -Inf
    Late_blastocyst.1_Cell.1
##
                               Late_blastocyst.1_Cell.2
##
                         -Inf
    Late_blastocyst.1_Cell.3
##
                               Late_blastocyst.1_Cell.4
##
                         -Inf
##
    Late_blastocyst.1_Cell.5
                               Late_blastocyst.1_Cell.6
##
                         -Inf
##
    Late_blastocyst.1_Cell.7
                               Late_blastocyst.1_Cell.8
##
                        -Inf
##
    Late_blastocyst.1_Cell.9 Late_blastocyst.1_Cell.10
##
                        -Inf
   Late_blastocyst.1_Cell.11 Late_blastocyst.1_Cell.12
##
##
                         -Inf
                                                    -Inf
    Late blastocyst.2 Cell.1 Late blastocyst.2 Cell.2
##
##
                  -0.3495575
                                              1.6981811
##
    Late_blastocyst.2_Cell.3
                              Late_blastocyst.2_Cell.4
##
                   0.6714127
##
    Late_blastocyst.2_Cell.5
                              Late_blastocyst.2_Cell.6
##
                         -Inf
##
    Late_blastocyst.2_Cell.7
                              Late_blastocyst.2_Cell.8
##
                         -Inf
##
    Late_blastocyst.2_Cell.9 Late_blastocyst.2_Cell.10
##
                         -Inf
##
    Late_blastocyst.3_Cell.1
                              Late_blastocyst.3_Cell.2
##
                         -Inf
##
    Late_blastocyst.3_Cell.3
                              Late_blastocyst.3_Cell.4
##
                         -Inf
##
    Late_blastocyst.3_Cell.5
                              Late_blastocyst.3_Cell.6
##
                    2.7084501
##
    Late_blastocyst.3_Cell.7
                              Late_blastocyst.3_Cell.8
##
                        -Inf
                                              2.7073166
print(pp)
         Oocyte.1 Oocyte.2 Oocyte.3 Zygote.1 Zygote.2
   [1,] 0.5312359 0.4510596 0.3740803 0.3927565 0.4492002
         Zygote.3 X2.cell_embryo.1_Cell.1
##
   [1,] 0.7291316
                                 0.4239975
##
        X2.cell_embryo.1_Cell.2 X2.cell_embryo.2_Cell.1
## [1,]
                      0.4793757
                                               0.3819854
##
        X2.cell_embryo.2_Cell.2 X2.cell_embryo.3_Cell.1
## [1,]
                      0.4468497
```

X2.cell_embryo.3_Cell.2 X4.cell_embryo.1_Cell.1

0.7291316

[1,]

0.7291316

0.7291316

```
X4.cell_embryo.1_Cell.2 X4.cell_embryo.1_Cell.3
##
##
   [1,]
                        0.176849
                                               0.7291316
##
        X4.cell_embryo.1_Cell.4 X4.cell_embryo.2_Cell.1
   [1,]
                      0.7291316
##
                                               0.7291316
##
        X4.cell_embryo.2_Cell.2 X4.cell_embryo.2_Cell.3
   [1,]
                      0.7291316
##
                                               0.4277359
##
        X4.cell_embryo.2_Cell.4 X4.cell_embryo.3_Cell.1
## [1,]
                        0.287862
                                                0.2658452
##
        X4.cell_embryo.3_Cell.2 X4.cell_embryo.3_Cell.3
##
   [1,]
                      0.3106824
                                               0.1951331
##
        X4.cell_embryo.3_Cell.4 X8.cell_embryo.1_Cell.1
   [1,]
##
                      0.1662384
                                               0.7291316
##
        X8.cell_embryo.1_Cell.2 X8.cell_embryo.1_Cell.3
   [1,]
##
                      0.2852823
                                               0.3765147
##
        X8.cell_embryo.1_Cell.4 X8.cell_embryo.2_Cell.1
##
   [1,]
                       0.3103037
                                               0.7291316
        X8.cell_embryo.2_Cell.2 X8.cell_embryo.2_Cell.3
##
##
   [1,]
                      0.7291316
                                               0.7291316
##
        X8.cell_embryo.2_Cell.4 X8.cell_embryo.2_Cell.5
##
   [1,]
                      0.7291316
                                               0.7291316
##
        X8.cell_embryo.2_Cell.6 X8.cell_embryo.2_Cell.7
##
  [1,]
                      0.7291316
                                               0.7291316
##
        X8.cell_embryo.2_Cell.8 X8.cell_embryo.3_Cell.1
   [1,]
##
                      0.7291316
                                               0.7291316
##
        X8.cell_embryo.3_Cell.2 X8.cell_embryo.3_Cell.3
##
   [1,]
                      0.7291316
                                               0.7291316
##
        X8.cell_embryo.3_Cell.4 X8.cell_embryo.3_Cell.5
##
   [1,]
                      0.7291316
                                               0.7291316
##
        X8.cell_embryo.3_Cell.6 X8.cell_embryo.3_Cell.7
## [1,]
                      0.7291316
                                               0.7291316
##
        X8.cell_embryo.3_Cell.8 Morulae.1_Cell.1
## [1,]
                      0.1430187
                                        0.7291316
##
        Morulae.1_Cell.2 Morulae.1_Cell.3 Morulae.1_Cell.4
##
   [1,]
               0.7291316
                                 0.7291316
                                                   0.7291316
        Morulae.1 Cell.5 Morulae.1 Cell.6 Morulae.1 Cell.7
##
##
                                 0.7291316
   [1,]
               0.7291316
                                                  0.7291316
##
        Morulae.1 Cell.8 Morulae.2 Cell.1 Morulae.2 Cell.2
##
  [1,]
               0.7291316
                                 0.7291316
                                                   0.7291316
##
        Morulae.2_Cell.3 Morulae.2_Cell.4 Morulae.2_Cell.5
##
   [1,]
               0.7291316
                                 0.7291316
                                                   0.7291316
##
        Morulae.2_Cell.6 Morulae.2_Cell.7 Morulae.2_Cell.8
   [1,]
##
               0.7291316
                                 0.7291316
                                                  0.7291316
##
        Late_blastocyst.1_Cell.1 Late_blastocyst.1_Cell.2
##
   [1,]
                        0.7291316
                                                  0.7291316
##
        Late_blastocyst.1_Cell.3 Late_blastocyst.1_Cell.4
## [1,]
                        0.7291316
                                                  0.7291316
##
        Late_blastocyst.1_Cell.5 Late_blastocyst.1_Cell.6
   [1,]
##
                        0.7291316
                                                  0.7291316
##
        Late_blastocyst.1_Cell.7 Late_blastocyst.1_Cell.8
##
   [1,]
                        0.7291316
                                                  0.7291316
##
        Late_blastocyst.1_Cell.9 Late_blastocyst.1_Cell.10
##
   [1,]
                       0.7291316
                                                   0.7291316
##
        Late_blastocyst.1_Cell.11 Late_blastocyst.1_Cell.12
## [1,]
                         0.7291316
                                                    0.7291316
```

```
##
        Late_blastocyst.2_Cell.1 Late_blastocyst.2_Cell.2
   [1,]
##
                       0.7291316
                                                 0.1516543
##
        Late_blastocyst.2_Cell.3 Late_blastocyst.2_Cell.4
##
   [1,]
                       0.3726822
                                                  0.7291316
##
        Late_blastocyst.2_Cell.5 Late_blastocyst.2_Cell.6
   [1,]
##
                       0.7291316
                                                 0.7291316
##
        Late_blastocyst.2_Cell.7 Late_blastocyst.2_Cell.8
## [1,]
                        0.7291316
                                                  0.7291316
##
        Late_blastocyst.2_Cell.9 Late_blastocyst.2_Cell.10
   [1,]
##
                       0.7291316
                                                  0.7291316
##
        Late_blastocyst.3_Cell.1 Late_blastocyst.3_Cell.2
   [1,]
##
                       0.7291316
                                                 0.7291316
##
        Late_blastocyst.3_Cell.3 Late_blastocyst.3_Cell.4
   [1,]
                                                 0.7291316
##
                       0.7291316
##
        Late_blastocyst.3_Cell.5 Late_blastocyst.3_Cell.6
## [1,]
                      0.04108083
                                                  0.7291316
##
        Late_blastocyst.3_Cell.7 Late_blastocyst.3_Cell.8
## [1,]
                       0.7291316
                                                0.04115071
```

print(cc)

```
##
                     Oocyte.1
                                                 Oocyte.2
##
                    0.000000
                                               0.3534698
##
                     Oocyte.3
                                                 Zygote.1
##
                    0.6657760
                                               0.5905606
##
                     Zygote.2
                                                 Zygote.3
##
                    0.3611648
                                               -0.3133418
     X2.cell_embryo.1_Cell.1
##
                                 X2.cell_embryo.1_Cell.2
##
                    0.4643627
                                               0.2342813
##
     X2.cell_embryo.2_Cell.1
                                 X2.cell_embryo.2_Cell.2
##
                    0.6339278
                                               0.3708737
##
     X2.cell_embryo.3_Cell.1
                                 X2.cell_embryo.3_Cell.2
##
                   -0.3523984
                                               -0.7052198
##
     X4.cell_embryo.1_Cell.1
                                 X4.cell_embryo.1_Cell.2
##
                         -Inf
                                                1.5526559
##
     X4.cell_embryo.1_Cell.3
                                 X4.cell embryo.1 Cell.4
##
                         -Inf
                                                     -Inf
##
     X4.cell_embryo.2_Cell.1
                                 X4.cell_embryo.2_Cell.2
##
                   -1.3586792
                                               -0.2943711
##
     X4.cell_embryo.2_Cell.3
                                 X4.cell_embryo.2_Cell.4
##
                    0.4491630
                                                1.0217312
##
     X4.cell_embryo.3_Cell.1
                                 X4.cell_embryo.3_Cell.2
##
                    1.1177611
                                               0.9250522
##
     X4.cell_embryo.3_Cell.3
                                 X4.cell_embryo.3_Cell.4
##
                    1.4548872
                                                1.6122340
##
     X8.cell_embryo.1_Cell.1
                                 X8.cell_embryo.1_Cell.2
##
                   -0.7635696
                                                1.0328285
##
     X8.cell_embryo.1_Cell.3
                                 X8.cell_embryo.1_Cell.4
##
                    0.6559644
                                                0.9266370
##
     X8.cell_embryo.2_Cell.1
                                 X8.cell_embryo.2_Cell.2
##
                         -Tnf
##
     X8.cell_embryo.2_Cell.3
                                 X8.cell_embryo.2_Cell.4
##
                         -Inf
##
     X8.cell_embryo.2_Cell.5
                                 X8.cell_embryo.2_Cell.6
```

```
##
                         -Inf
                                                     -Inf
##
     X8.cell_embryo.2_Cell.7
                                X8.cell_embryo.2_Cell.8
##
##
     X8.cell_embryo.3_Cell.1
                                X8.cell_embryo.3_Cell.2
##
                         -Inf
##
     X8.cell_embryo.3_Cell.3
                                X8.cell embryo.3 Cell.4
##
                         -Inf
##
     X8.cell_embryo.3_Cell.5
                                X8.cell_embryo.3_Cell.6
##
                         -Inf
                                                     -Inf
##
                                X8.cell_embryo.3_Cell.8
     X8.cell_embryo.3_Cell.7
##
                         -Inf
                                               1.7516317
##
            Morulae.1_Cell.1
                                        Morulae.1_Cell.2
##
                         -Inf
                                                    -Inf
                                        Morulae.1_Cell.4
##
            Morulae.1_Cell.3
##
                         -Inf
                                              -0.2930297
##
            Morulae.1_Cell.5
                                        Morulae.1_Cell.6
##
                         -Inf
##
            Morulae.1_Cell.7
                                        Morulae.1 Cell.8
##
                         -Inf
                                                    -Inf
##
            Morulae.2 Cell.1
                                        Morulae.2 Cell.2
##
                         -Inf
                                                    -Inf
##
            Morulae.2 Cell.3
                                        Morulae.2 Cell.4
##
                                                    -Inf
                         -Inf
            Morulae.2 Cell.5
                                        Morulae.2 Cell.6
##
##
                         -Inf
                                              -0.9702191
##
            Morulae.2 Cell.7
                                        Morulae.2_Cell.8
##
                  -0.9597203
                                                    -Inf
##
    Late_blastocyst.1_Cell.1
                               Late_blastocyst.1_Cell.2
##
                         -Inf
    {\tt Late\_blastocyst.1\_Cell.3}
##
                               Late_blastocyst.1_Cell.4
##
                         -Inf
##
    Late_blastocyst.1_Cell.5
                               Late_blastocyst.1_Cell.6
##
                         -Inf
##
    Late_blastocyst.1_Cell.7
                               Late_blastocyst.1_Cell.8
##
                         -Inf
##
    Late_blastocyst.1_Cell.9 Late_blastocyst.1_Cell.10
##
##
   Late_blastocyst.1_Cell.11 Late_blastocyst.1_Cell.12
##
                         -Inf
##
                              Late_blastocyst.2_Cell.2
    Late_blastocyst.2_Cell.1
##
                  -0.3495575
                                               1.6981811
##
    Late_blastocyst.2_Cell.3
                               Late_blastocyst.2_Cell.4
##
                    0.6714127
##
    Late_blastocyst.2_Cell.5
                               Late_blastocyst.2_Cell.6
##
                         -Inf
##
    Late_blastocyst.2_Cell.7
                               Late_blastocyst.2_Cell.8
##
                         -Inf
    Late_blastocyst.2_Cell.9 Late_blastocyst.2_Cell.10
##
##
                         -Inf
                                                    -Inf
##
    Late_blastocyst.3_Cell.1
                               Late_blastocyst.3_Cell.2
##
                         -Inf
    Late_blastocyst.3_Cell.3
##
                               Late blastocyst.3 Cell.4
##
                         -Inf
    Late_blastocyst.3_Cell.5 Late_blastocyst.3_Cell.6
```

```
##
                    2.7084501
                                                     -Inf
##
    Late_blastocyst.3_Cell.7 Late_blastocyst.3_Cell.8
                                               2.7073166
##
                         -Inf
print(dd)
## NULL
i <- 5
x <- data0[i, ]
Zcut0 <- 0
best <- GetBestK(x = x, n = 1000, q = Zcut0, err = 1e-10)
pp <- calculate_prob_sep_Zcut(x, Zcut0, best$result[, 1], best$result[, 2], best$result[, 3])
cc <- discretization_method_1_LLR_mean(x, best$result, pp, LLR_cut = 0.1)</pre>
## [1] "Only one class"
dd <- Build_R_matrix(cc, Zcut0, best$result[, 2], rownames(data0)[i])</pre>
print(x)
##
                     Oocyte.1
                                                Oocyte.2
##
                   -0.3871342
                                               0.2949059
##
                     Oocyte.3
                                                Zygote.1
##
                    0.7537718
                                               1.7446679
##
                     Zygote.2
                                                Zygote.3
##
                    1.5871923
                                               1.7313015
##
     X2.cell_embryo.1_Cell.1
                                 X2.cell_embryo.1_Cell.2
##
                    1.4611699
                                               1.4011830
##
     X2.cell_embryo.2_Cell.1
                                 X2.cell_embryo.2_Cell.2
##
                    1.4731599
                                                1.5475625
##
     X2.cell_embryo.3_Cell.1
                                X2.cell_embryo.3_Cell.2
##
                    1.0501221
                                               0.8742180
##
     X4.cell_embryo.1_Cell.1
                                 X4.cell_embryo.1_Cell.2
##
                    1.4060970
                                              -1.8325815
##
     X4.cell_embryo.1_Cell.3
                                 X4.cell_embryo.1_Cell.4
##
                    0.1856493
                                                     -Inf
##
     X4.cell_embryo.2_Cell.1
                                 X4.cell_embryo.2_Cell.2
##
                    1.1413524
                                                     -Inf
##
     X4.cell_embryo.2_Cell.3
                                 X4.cell_embryo.2_Cell.4
##
                    2.3799164
                                              -0.9288695
##
     X4.cell_embryo.3_Cell.1
                                 X4.cell_embryo.3_Cell.2
##
                    1.5411591
                                               1.7523254
##
     X4.cell_embryo.3_Cell.3
                                X4.cell_embryo.3_Cell.4
##
                    1.5091755
                                               1.7288197
##
     X8.cell_embryo.1_Cell.1
                                 X8.cell_embryo.1_Cell.2
##
                    3.5036336
                                               3.6047095
##
     X8.cell_embryo.1_Cell.3
                                 X8.cell_embryo.1_Cell.4
##
                    2.8523237
                                               3.7815954
##
     X8.cell_embryo.2_Cell.1
                                 X8.cell_embryo.2_Cell.2
##
                    2.7822296
                                               1.7838953
```

```
##
     X8.cell_embryo.2_Cell.3
                                X8.cell_embryo.2_Cell.4
##
                    2.0550208
                                               3.6057967
     X8.cell_embryo.2_Cell.5
                                X8.cell_embryo.2_Cell.6
##
##
                    2.8845213
                                               2.2490788
##
     X8.cell_embryo.2_Cell.7
                                X8.cell_embryo.2_Cell.8
##
                    2.2758304
                                               2.9764475
     X8.cell_embryo.3_Cell.1
##
                                X8.cell_embryo.3_Cell.2
##
                    3.2518463
                                               3.4940196
##
     X8.cell_embryo.3_Cell.3
                                X8.cell_embryo.3_Cell.4
##
                    4.0421210
                                               3.8096580
##
     X8.cell_embryo.3_Cell.5
                                X8.cell_embryo.3_Cell.6
##
                    3.6084554
                                               2.3880286
     X8.cell_embryo.3_Cell.7
                                X8.cell_embryo.3_Cell.8
##
                    2.9516759
##
                                               4.0716217
##
            Morulae.1_Cell.1
                                        Morulae.1_Cell.2
##
                    3.9599366
                                               3.4106195
##
            Morulae.1_Cell.3
                                        Morulae.1_Cell.4
##
                   -0.1086994
                                               3.2040464
##
            Morulae.1_Cell.5
                                        Morulae.1_Cell.6
##
                    3.0037004
                                               4.1114475
##
            Morulae.1_Cell.7
                                        Morulae.1_Cell.8
##
                    3.7993021
                                              -0.8141855
##
            Morulae.2_Cell.1
                                        Morulae.2_Cell.2
##
                    3.0061775
                                               4.1204670
##
            Morulae.2 Cell.3
                                        Morulae.2 Cell.4
##
                    3.9427656
                                               3.4452781
##
                                        Morulae.2_Cell.6
            Morulae.2_Cell.5
##
                    3.0906333
                                               3.1098642
##
            Morulae.2_Cell.7
                                        Morulae.2_Cell.8
##
                    3.8282935
                                               4.1413236
##
    Late_blastocyst.1_Cell.1
                               Late_blastocyst.1_Cell.2
##
                    0.3133498
                                               2.3487050
##
    Late_blastocyst.1_Cell.3
                               Late_blastocyst.1_Cell.4
##
                    3.4520489
                                               2.5867861
    Late_blastocyst.1_Cell.5
##
                               Late blastocyst.1 Cell.6
##
                    4.2983325
                                               3.2938349
##
    Late blastocyst.1 Cell.7
                               Late blastocyst.1 Cell.8
##
                    2.2683042
                                               1.4092782
##
    Late_blastocyst.1_Cell.9 Late_blastocyst.1_Cell.10
##
                         -Inf
                                               3.0728322
##
   Late blastocyst.1 Cell.11 Late blastocyst.1 Cell.12
##
                    1.8878269
                                               2.3270826
##
    Late_blastocyst.2_Cell.1
                               Late_blastocyst.2_Cell.2
##
                    2.2086040
                                               2.9670756
##
    Late_blastocyst.2_Cell.3
                               Late_blastocyst.2_Cell.4
##
                    3.2008340
                                              -0.3768777
##
    Late_blastocyst.2_Cell.5
                               Late_blastocyst.2_Cell.6
##
                   -0.2256467
                                               2.8136107
##
    Late_blastocyst.2_Cell.7
                               Late_blastocyst.2_Cell.8
##
                   -0.8462984
                                               2.6758713
##
    Late_blastocyst.2_Cell.9 Late_blastocyst.2_Cell.10
##
                    1.4768204
                                               2.1813210
##
    Late_blastocyst.3_Cell.1
                              Late_blastocyst.3_Cell.2
##
                    2.0725428
                                               1.3790180
```

```
Late_blastocyst.3_Cell.3 Late_blastocyst.3_Cell.4
##
                   2.3051817
                                              2.4216118
    Late_blastocyst.3_Cell.5
                              Late blastocyst.3 Cell.6
##
##
                   3.4375293
                                             -1.9661129
##
    Late blastocyst.3 Cell.7
                              Late_blastocyst.3_Cell.8
                  -0.9314044
                                              1.9226415
##
```

print(pp)

```
Oocyte.1 Oocyte.2 Oocyte.3 Zygote.1 Zygote.2
   [1,] 0.08272947 0.3284781 0.4512065 0.650595 0.6320212
##
         Zygote.3 X2.cell_embryo.1_Cell.1
   [1,] 0.6492757
                                 0.6126515
##
        X2.cell_embryo.1_Cell.2 X2.cell_embryo.2_Cell.1
   [1,]
                      0.6021446
                                               0.6146554
##
        X2.cell_embryo.2_Cell.2 X2.cell_embryo.3_Cell.1
##
   [1,]
                      0.6263418
                                               0.5269793
##
        X2.cell_embryo.3_Cell.2 X4.cell_embryo.1_Cell.1
## [1,]
                      0.4828621
                                               0.6030348
##
        X4.cell_embryo.1_Cell.2 X4.cell_embryo.1_Cell.3
## [1,]
                     0.08272947
                                               0.3003883
##
        X4.cell_embryo.1_Cell.4 X4.cell_embryo.2_Cell.1
##
   [1,]
                     0.08272947
                                               0.5484437
##
        X4.cell_embryo.2_Cell.2 X4.cell_embryo.2_Cell.3
##
   [1,]
                     0.08272947
                                               0.6537385
##
        X4.cell_embryo.2_Cell.4 X4.cell_embryo.3_Cell.1
## [1,]
                     0.08272947
                                               0.6253879
##
        X4.cell_embryo.3_Cell.2 X4.cell_embryo.3_Cell.3
## [1,]
                      0.6513288
                                               0.6204757
##
        X4.cell_embryo.3_Cell.4 X8.cell_embryo.1_Cell.1
##
   [1,]
                      0.6490253
                                               0.4247675
##
        X8.cell_embryo.1_Cell.2 X8.cell_embryo.1_Cell.3
##
   [1,]
                      0.3975103
                                               0.5839943
##
        X8.cell_embryo.1_Cell.4 X8.cell_embryo.2_Cell.1
## [1,]
                      0.3501002
                                               0.5975892
##
        X8.cell_embryo.2_Cell.2 X8.cell_embryo.2_Cell.3
  [1,]
##
                      0.6541823
                                               0.6669216
        X8.cell_embryo.2_Cell.4 X8.cell_embryo.2_Cell.5
   [1,]
##
                       0.397217
                                               0.5774304
        X8.cell_embryo.2_Cell.6 X8.cell_embryo.2_Cell.7
##
## [1,]
                      0.6627484
                                               0.6613057
##
        X8.cell_embryo.2_Cell.8 X8.cell_embryo.3_Cell.1
## [1,]
                      0.5576759
                                               0.4912675
##
        X8.cell_embryo.3_Cell.2 X8.cell_embryo.3_Cell.3
   [1,]
##
                      0.4273543
                                               0.2831046
##
        X8.cell_embryo.3_Cell.4 X8.cell_embryo.3_Cell.5
##
   [1,]
                      0.3426781
                                               0.3964998
        X8.cell_embryo.3_Cell.6 X8.cell_embryo.3_Cell.7
##
## [1,]
                      0.6530201
##
        X8.cell_embryo.3_Cell.8 Morulae.1_Cell.1
## [1,]
                      0.2758512
                                        0.3037122
##
        Morulae.1_Cell.2 Morulae.1_Cell.3 Morulae.1_Cell.4
  [1,]
                               0.08272947
               0.4496895
##
        Morulae.1_Cell.5 Morulae.1_Cell.6 Morulae.1_Cell.7
```

```
## [1,]
               0.5515517
                                 0.2661895
                                                   0.3454127
##
        Morulae.1_Cell.8 Morulae.2_Cell.1 Morulae.2_Cell.2
              0.08272947
##
   [1,]
                                 0.5509893
                                                   0.2640229
##
        Morulae.2_Cell.3 Morulae.2_Cell.4 Morulae.2_Cell.5
##
   [1,]
               0.3080868
                                 0.4404348
                                                   0.5312876
##
        Morulae.2 Cell.6 Morulae.2 Cell.7 Morulae.2 Cell.8
  [1,]
##
               0.5266672
                                 0.3377711
                                                   0.2590444
##
        Late_blastocyst.1_Cell.1 Late_blastocyst.1_Cell.2
##
   [1,]
                         0.333298
                                                  0.6563305
##
        Late_blastocyst.1_Cell.3 Late_blastocyst.1_Cell.4
##
   [1,]
                        0.4386218
                                                  0.6298768
##
        Late_blastocyst.1_Cell.5 Late_blastocyst.1_Cell.6
##
   [1,]
                        0.2230686
                                                  0.4804357
##
        Late_blastocyst.1_Cell.7 Late_blastocyst.1_Cell.8
##
   [1,]
                        0.6617326
                                                  0.6036083
##
        Late_blastocyst.1_Cell.9 Late_blastocyst.1_Cell.10
##
   [1,]
                      0.08272947
                                                   0.5355221
##
        Late_blastocyst.1_Cell.11 Late_blastocyst.1_Cell.12
   [1,]
##
                         0.6615891
                                                    0.6579649
##
        Late_blastocyst.2_Cell.1 Late_blastocyst.2_Cell.2
##
   [1,]
                         0.664535
                                                  0.5597549
##
        Late_blastocyst.2_Cell.3 Late_blastocyst.2_Cell.4
  [1,]
##
                        0.5042238
                                                 0.08272947
##
        Late_blastocyst.2_Cell.5 Late_blastocyst.2_Cell.6
##
   [1,]
                      0.08272947
                                                  0.5916239
##
        Late_blastocyst.2_Cell.7 Late_blastocyst.2_Cell.8
   [1,]
##
                      0.08272947
                                                  0.6162464
##
        Late_blastocyst.2_Cell.9 Late_blastocyst.2_Cell.10
##
   [1,]
                        0.6152607
                                                   0.6654682
##
        Late_blastocyst.3_Cell.1 Late_blastocyst.3_Cell.2
##
   [1,]
                        0.6670025
                                                  0.5980658
##
        Late_blastocyst.3_Cell.3 Late_blastocyst.3_Cell.4
##
   [1,]
                        0.6594847
                                                  0.6498522
##
        Late_blastocyst.3_Cell.5 Late_blastocyst.3_Cell.6
##
   [1,]
                        0.4425078
                                                 0.08272947
##
        Late_blastocyst.3_Cell.7 Late_blastocyst.3_Cell.8
## [1,]
                      0.08272947
                                                  0.6633757
```

print(cc)

```
##
                     Oocyte.1
                                                 Oocyte.2
##
                   -0.3871342
                                                0.2949059
##
                     Oocyte.3
                                                 Zygote.1
##
                    0.7537718
                                                1.7446679
##
                     Zygote.2
                                                 Zygote.3
##
                    1.5871923
                                                1.7313015
##
                                 X2.cell_embryo.1_Cell.2
     X2.cell_embryo.1_Cell.1
##
                    1.4611699
                                                1.4011830
     X2.cell_embryo.2_Cell.1
##
                                 X2.cell_embryo.2_Cell.2
##
                    1.4731599
                                                1.5475625
##
     X2.cell_embryo.3_Cell.1
                                 X2.cell_embryo.3_Cell.2
##
                    1.0501221
                                                0.8742180
##
     X4.cell_embryo.1_Cell.1
                                 X4.cell_embryo.1_Cell.2
##
                    1.4060970
                                               -1.8325815
```

## ##	X4.cell_embryo.1_Cell.3 0.1856493	X4.cell_embryo.1_Cell.4 -Inf
##	X4.cell_embryo.2_Cell.1	X4.cell_embryo.2_Cell.2
##	1.1413524	-Inf
##	X4.cell_embryo.2_Cell.3	X4.cell_embryo.2_Cell.4
##	2.3799164	-0.9288695
##	X4.cell_embryo.3_Cell.1	X4.cell_embryo.3_Cell.2
##	1.5411591	1.7523254
##	X4.cell_embryo.3_Cell.3	X4.cell_embryo.3_Cell.4
##	1.5091755	1.7288197
##	X8.cell_embryo.1_Cell.1	X8.cell_embryo.1_Cell.2
##	3.5036336	3.6047095
##	X8.cell_embryo.1_Cell.3	X8.cell_embryo.1_Cell.4
	_ • • • —	_ • -
##	2.8523237	3.7815954
##	X8.cell_embryo.2_Cell.1	X8.cell_embryo.2_Cell.2
##	2.7822296	1.7838953
##	X8.cell_embryo.2_Cell.3	X8.cell_embryo.2_Cell.4
##	2.0550208	3.6057967
##	X8.cell_embryo.2_Cell.5	X8.cell_embryo.2_Cell.6
##	2.8845213	2.2490788
##	X8.cell_embryo.2_Cell.7	X8.cell_embryo.2_Cell.8
##	2.2758304	2.9764475
##	X8.cell_embryo.3_Cell.1	X8.cell_embryo.3_Cell.2
##	3.2518463	3.4940196
##	X8.cell_embryo.3_Cell.3	X8.cell_embryo.3_Cell.4
##	4.0421210	3.8096580
##	X8.cell_embryo.3_Cell.5	X8.cell_embryo.3_Cell.6
##	3.6084554	2.3880286
##	X8.cell_embryo.3_Cell.7	X8.cell_embryo.3_Cell.8
##	2.9516759	4.0716217
##	Morulae.1_Cell.1	Morulae.1_Cell.2
##	3.9599366	3.4106195
##	Morulae.1_Cell.3	Morulae.1_Cell.4
##	-0.1086994	3.2040464
##	Morulae.1_Cell.5	Morulae.1_Cell.6
##	3.0037004	4.1114475
##	Morulae.1 Cell.7	Morulae.1 Cell.8
	3.7993021	-0.8141855
##		
##	Morulae.2_Cell.1	Morulae.2_Cell.2
##	3.0061775	4.1204670
##	Morulae.2_Cell.3	Morulae.2_Cell.4
##	3.9427656	3.4452781
##	Morulae.2_Cell.5	Morulae.2_Cell.6
##	3.0906333	3.1098642
##	Morulae.2_Cell.7	Morulae.2_Cell.8
##	3.8282935	4.1413236
##	Late_blastocyst.1_Cell.1	Late_blastocyst.1_Cell.2
##	0.3133498	2.3487050
##	Late_blastocyst.1_Cell.3	Late_blastocyst.1_Cell.4
##	3.4520489	2.5867861
##	Late_blastocyst.1_Cell.5	Late_blastocyst.1_Cell.6
##	4.2983325	3.2938349
##	Late_blastocyst.1_Cell.7	Late_blastocyst.1_Cell.8
##	2.2683042	1.4092782
	2.200012	1.1002/02

```
Late_blastocyst.1_Cell.9 Late_blastocyst.1_Cell.10
##
##
                         -Inf
                                              3.0728322
##
  Late_blastocyst.1_Cell.11 Late_blastocyst.1_Cell.12
##
                   1.8878269
                                              2.3270826
##
    Late_blastocyst.2_Cell.1
                              Late_blastocyst.2_Cell.2
##
                   2.2086040
                                              2.9670756
##
   Late blastocyst.2 Cell.3
                              Late_blastocyst.2_Cell.4
##
                   3.2008340
                                             -0.3768777
##
    Late_blastocyst.2_Cell.5
                              Late_blastocyst.2_Cell.6
##
                  -0.2256467
                                              2.8136107
##
    Late_blastocyst.2_Cell.7
                              Late_blastocyst.2_Cell.8
##
                  -0.8462984
                                              2.6758713
    Late_blastocyst.2_Cell.9 Late_blastocyst.2_Cell.10
##
                   1.4768204
##
                                              2.1813210
##
    Late_blastocyst.3_Cell.1
                              Late_blastocyst.3_Cell.2
##
                   2.0725428
                                               1.3790180
##
    Late_blastocyst.3_Cell.3
                              Late_blastocyst.3_Cell.4
##
                   2.3051817
                                              2.4216118
##
    Late_blastocyst.3_Cell.5
                              Late_blastocyst.3_Cell.6
##
                   3.4375293
                                             -1.9661129
##
    Late_blastocyst.3_Cell.7
                              Late_blastocyst.3_Cell.8
##
                  -0.9314044
                                              1.9226415
print(dd)
```

NULL

(iv) directly apply qubic from the QUBIC package

We are trying save the result to a file first.

```
WriteQubicInput <- function(file.name, data0, genes, q = 0, err = 1e-10) {
  cat("o", colnames(data0), "\n", file = file.name)
  for (i in genes) {
    cat(i, colnames(data0), "\n", file = "progress")
    x <- data0[i, ]
    Zcut0 <- q
    best <- GetBestK(x = x, n = 1000, q = Zcut0, err = 1e-10)
    if (best$k == 0) {
      next
    }
    pp <- calculate_prob_sep_Zcut(x, Zcut0, best$result[, 1], best$result[, 2], best$result[, 3])</pre>
    cc <- discretization_method_1_LLR_mean(x, best$result, pp, LLR_cut = 0.1)</pre>
    dd <- Build_R_matrix(cc, Zcut0, best$result[, 2], rownames(data0)[i])</pre>
    write.table(dd, file = file.name, col.names = FALSE, append = TRUE, quote = FALSE)
  }
}
system.time(WriteQubicInput("qubic_input_head30", data0, head(selected.genes, 30)))
## [1] "Only one class"
## [1] "Only one class"
```

```
## [1] "Only one class"
##
      user system elapsed
##
      6.00
             0.07
                      6.17
This may be slow...
print(length(selected.genes))
## [1] 14542
qubic.file = "qubic input"
if (!file.exists(qubic.file)) {
  WriteQubicInput(qubic.file, data0, selected.genes)
}
```

It is the time to read all the data back.

```
qubic.input <- as.matrix(read.table(qubic.file, row.names = 1, header = TRUE))</pre>
```

Run QUBIC (Zhang et al. 2017), need several minute.

```
library(QUBIC)
if (!file.exists("res.RData")) {
  res <- qubiclust_d(qubic.input)</pre>
  save(res, file="res.RData")
} else {
  load("res.RData")
}
```

(v) results summary

```
##
  An object of class Biclust
##
## call:
##
   NULL
## Number of Clusters found: 100
## First 5 Cluster sizes:
                       BC 1 BC 2 BC 3 BC 4 BC 5
## Number of Rows:
                        934
                             931
                                  921
                                        959
## Number of Columns:
                         73
                              73
                                   73
                                         70
biclust::summary(res)
##
  An object of class Biclust
##
## call:
##
   NULL
## Number of Clusters found: 100
##
## Cluster sizes:
##
                       BC 1 BC 2 BC 3 BC 4 BC 5 BC 6 BC 7 BC 8
## Number of Rows:
                        934 931
                                  921
                                        959
                                             941
                                                  911
## Number of Columns:
                         73
                              73
                                   73
                                         70
                                              71
                                                    73
                                                         73
                       BC 9 BC 10 BC 11 BC 12 BC 13 BC 14 BC 15
## Number of Rows:
                        882
                                     902
                                           966
                                                  885
                                                        872
                                                              872
                              917
## Number of Columns:
                         75
                               72
                                      73
                                            68
                                                   74
                                                         75
                                                               75
##
                       BC 16 BC 17 BC 18 BC 19 BC 20 BC 21
## Number of Rows:
                         894
                               892
                                      891
                                            910
                                                   897
  Number of Columns:
                          73
                                73
                                       73
                                             71
                                                    72
                                                          71
                       BC 22 BC 23 BC 24 BC 25 BC 26 BC 27
                         870
                                      905
## Number of Rows:
                               870
                                            866
                                                   850
## Number of Columns:
                          74
                                74
                                       71
                                             74
                                                    75
                                                          75
##
                       BC 28 BC 29 BC 30 BC 31 BC 32 BC 33
## Number of Rows:
                         846
                               845
                                      832
                                            832
                                                   832
## Number of Columns:
                          75
                                74
                                             75
                                                    75
                                       75
                       BC 34 BC 35 BC 36 BC 37 BC 38 BC 39
## Number of Rows:
                         831
                               831
                                      831
                                            831
                                                  831
                                                         831
## Number of Columns:
                          75
                                75
                                       75
                                             75
                                                    75
                                                          75
##
                       BC 40 BC 41 BC 42 BC 43 BC 44 BC 45
## Number of Rows:
                         831
                               838
                                      826
                                            826
                                                  826
## Number of Columns:
                          75
                                74
                                       75
                                             75
                                                    75
                                                          75
##
                       BC 46 BC 47 BC 48 BC 49 BC 50 BC 51
## Number of Rows:
                         832
                               831
                                      863
                                            808
                                                   814
                                                         792
## Number of Columns:
                          74
                                74
                                       71
                                             75
                                                    73
                                                          75
```

```
##
                        BC 52 BC 53 BC 54 BC 55 BC 56 BC 57
## Number of Rows:
                          792
                                792
                                       792
                                             792
                                                    792
                                                           792
## Number of Columns:
                           75
                                  75
                                        75
                                              75
                                                     75
                                                            75
##
                       BC 58 BC 59 BC 60 BC 61 BC 62 BC 63
##
  Number of Rows:
                          792
                                792
                                       792
                                             792
                                                    792
                                                           792
##
  Number of Columns:
                                 75
                                        75
                                              75
                                                     75
                           75
                                                            75
##
                       BC 64 BC 65 BC 66 BC 67 BC 68 BC 69
                          792
                                792
                                       792
                                             792
                                                    792
## Number of Rows:
                                                           792
##
   Number of Columns:
                           75
                                  75
                                        75
                                               75
                                                     75
                                                            75
##
                        BC 70 BC 71 BC 72 BC 73 BC 74 BC 75
##
  Number of Rows:
                          792
                                792
                                       792
                                             792
                                                    792
                                                           792
                                 75
                                        75
                                              75
                                                            75
##
   Number of Columns:
                           75
                                                     75
                        BC 76 BC 77 BC 78 BC 79 BC 80 BC 81
##
## Number of Rows:
                          792
                                792
                                       792
                                             792
                                                    792
                                                           792
##
  Number of Columns:
                           75
                                  75
                                        75
                                               75
                                                     75
                                                            75
##
                        BC 82 BC 83 BC 84 BC 85 BC 86 BC 87
  Number of Rows:
                          792
                                792
                                       792
                                              792
                                                    792
                                                           792
##
   Number of Columns:
                           75
                                  75
                                        75
                                              75
                                                     75
                                                            75
##
                       BC 88 BC 89 BC 90 BC 91 BC 92 BC 93
## Number of Rows:
                          792
                                792
                                       792
                                             791
                                                    792
                                                           792
##
  Number of Columns:
                           75
                                 75
                                        75
                                              75
                                                     74
                                                            74
##
                        BC 94 BC 95 BC 96 BC 97 BC 98 BC 99
## Number of Rows:
                          792
                                792
                                       739
                                              725
                                                    699
                                                           683
##
  Number of Columns:
                           74
                                  74
                                        77
                                              76
                                                     78
                                                            77
##
                        BC 100
## Number of Rows:
                           668
## Number of Columns:
                            78
```

References

Zhang, Yu, Juan Xie, Jinyu Yang, Anne Fennell, Chi Zhang, and Qin Ma. 2017. "QUBIC: A Bioconductor Package for Qualitative Biclustering Analysis of Gene Co- Expression Data." *Bioinformatics* 33 (3): 450–52. https://doi.org/10.1093/bioinformatics/btw635.