Exact distribution of excursions height

Sébastien Déjean - Charly Marti - Sabine Mercier - Sebastian Simon - David Robelin 2025-02-24

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Introduction

The goal of this part of the package is to calculate the theoretical probability that the i-th excursion reach the threshold score a, for a Markov chain with a known transition matrix and a given score function. The main function for this objective is proba_theoretical_ith_excursion_markov().

This section of the local Score package is the direct implementation of the pre publication: "Exact distribution of excursion heights of the Lindley process in a Markovian model" written by Carlos Cortés Rojas, Simona Grusea and Sabine Mercier.

Computation method

The main goal of this function is to calculate the probability that the first excursion of the Lindley sequence associated to the studied sequence is greater or equal to a, conditionally to α a potential beginning of this sequence. It also computes the transition probability matrix of the beginnings of excursions: matrix_M.

The product of this matrix, elevated to the power i-1, with the first vector of the probabilities conditionally to α , gives the probabilities conditionally to α for the *i*-th excursion. To return the global probability, the function multiplies the conditional vector with the distribution of the first letter of the sequence.

Definition of a mathematical excursion

The number of an excursion is given by the mathematical definition of Karlin and Altschul (1990). Its corresponds to the number of record times of the Lindley sequence associated to the score sequence $(X_k)_{1 \le k \le n}$ and recursively defined as follows:

$$T_0 := 0$$
 and $T_{(k+1)} := \inf\{i \ge T_k + 1, \sum_{j=T_k+1}^i X_j < 0\}$.

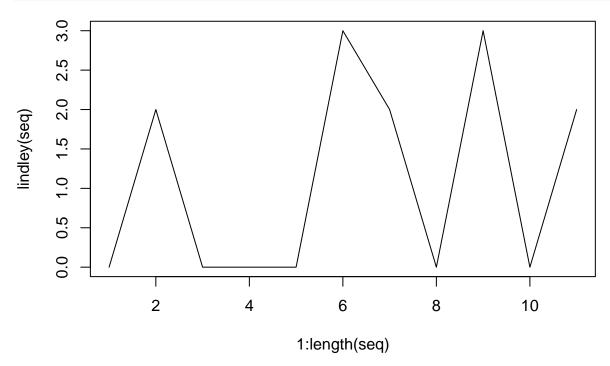
Please note that the sum $\sum_{j=T_k+1}^{i} X_j$ must be non positive. If it is equal to zero, it will mathematically be still the same Lindley excursion.

$$(A_k)_k$$
 with $T_{(i-1)} + 1 \le k < T_i$

is called the i-th positive excursion of the sequence in the term of Karlin and Altschul. This mathematical definition includes the index with a Lindey score equal to 0, whereas visually we "see" the excursion beginning at the following index.

The mathematical number of an excursion must be distinguished from the one of the corresponding visible excursion because of the possible appearance of flat excursions. See below for an example.

```
seq <- c(-1,2,-3,-4,-1,3,-1,-2,3,-4,2)
lindley(seq)
#> [1] 0 2 0 0 0 3 2 0 3 0 2
plot(1:length(seq), lindley(seq), type = 'l')
```



```
localScoreC(seq)
#> $localScore
#> value begin end
#> 3 6 6
#>
#> $suboptimalSegmentScores
```

```
value begin end
         2
               2
         3
               6
                   6
#> 3
         3
               9
                   9
#> 4
         2
              11
                  11
#>
#> $RecordTime
#> [1] 0 1 3 4 5 10
```

We can "see" three positive excursions, with top at index $\{2; 6; 9\}$; plus one last not achieved at index $\{11\}$. Their exists also five flats excursions: at index $\{1; 3; 4; 5; 10\}$. Note also that the Lindley excursion 6-10 reach 0 at index 8 without going in non positive values, so it is still the same excursion.

In summary, the excursions list is: $\{1; 2-3; 4; 5; 6-10; 11\}$. The following function gives the record time of the mathematical excursion that must be used to compute the statistical significance. Note that we choose to omit the conventional first record time which is always 0.

```
recordTimes(seq)
#> [1] 1 3 4 5 10
```

So the sub optimal excursion values given by \$suboptimalSegmentScores 2 3 3 2 are achieved by the excursion number 2, 5 (twice) and 6. At this point, we wish to lighten a difference between the notion of "excursion" defined mathematically by Karlin and Dembo (1992), and the results given in \$suboptimalSegmentScores. The segment [6-10] is an excursion in mathematical sense as the cumulative process starting at position 7 never hit a negative value before the 10th position. Nevertheless this process hit a null value indicating that this single excursion contain at least two positive score segments (here: at position 6 and 9). We chose to indicate in \$suboptimalSegmentScores all positive consecutive sub-sequences, as it can be more meaningful when applied on real data.

To calculate the p-value of a given excursion, we have to know which number of excursion it is in sequential order. Notice that it will not bring any problem of making a mistake in the number of the excursion for a number exceeding 10, and a small difference for a lower number, as the sequence composed by the begin-component of each excursion, which is also a Markov chain, can reach the stationary distribution. For the first excursions, it is far more easy to recover the record time of the corresponding mathematical excursion in the list.

Toy examples

As the function calculates a theoretical probability, we don't need a sequence of scores but the transition matrix of the markov chain: Λ . It also requires a score function with: integer scores, a negative expectation and at least one possible positive score. We also need θ , an alphabet (can contains numbers) with unique values; and i: the rank of the excursion on which we calculate the probabilities. The optional parameters are epsilon and prob0. epsilon is a threshold for the computation of matrix M and prob0 is the probability distribution of the first letter of the sequence. Obviously, theta, the score function and prob0 should have the same length p, and the dimension of lambda should be (p,p).

In this example, we want to know the probability that the second excursion reach a score greater or equal than 5 with the matrix Λ and a score function of (-1,1), given that the first score of the sequence have 50 percent chances to be -1.

In this following example, we see that this is not a problem to have missing score values in the score function, and that theta can contain numbers. Initial value of epsilon is 1e-16, and if we don't precise the value of prob0, the function compute the stationary distribution of Λ , and use it for the distribution of the first letter of the sequence.

In the following example, the function takes longer as we increased the number of scores (complexity of $O(length(\theta)^3)$), but here with 20 scores, the computational time is acceptable.

```
transition_matrix <- matrix(runif(400, min = 0, max = 1), nrow = 20)</pre>
transition_matrix <- t(apply(transition_matrix, 1, function(x) x/sum(x)))</pre>
theta <- letters[1:20]
score_f \leftarrow c(-3,-1,2,-3,-1,2,-3,-1,2,-1,
              -3,-1,2,-3,-1,2,-3,-1,2,-1
sum(stationary_distribution(transition_matrix)*score_f) #score expectation (stationary)
#> [1] -0.5776208
system.time(pv1 <- proba_theoretical_ith_excursion_markov(a=5, theta, transition_matrix, score_f, i = 4
#> utilisateur
                    système
                                 écoulé
#>
         4.724
                      0.001
                                  4.726
pv1$proba_q_i_geq_a
#> [1] 0.07892013
```

A study case

Let us consider the protein Seq1093 of 1093 amino acid proposed in the package localScore which corresponds to the Q60519.fasta in UniProt Data base.

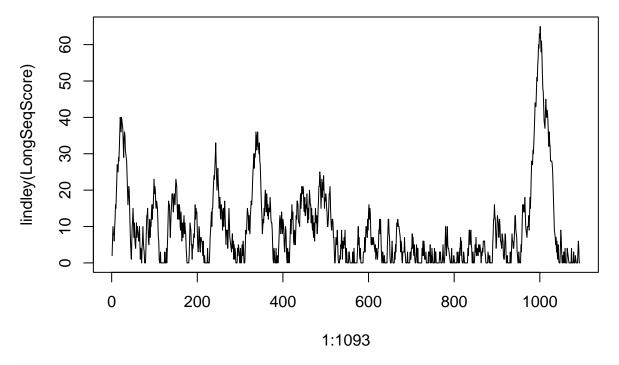
Optimal excursion, the local score

Using the hydrophobic score scale called HydroScore, we compute the local score, corresponding to the height of the highest excursion.

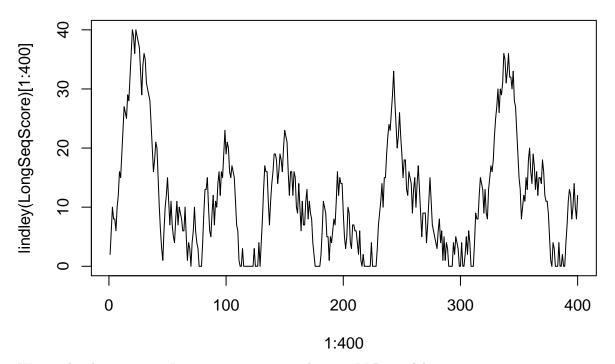
```
library(localScore)
data("Seq1093")
Seq1093
#> [1] "MVVPGPLALSLLLSSLTLLVSHLSSSQDIASESSSEQQMCTRREHPIVAFEDLKPWVFNFTYPGVRDFSQLALDPSRNQLIVGARNYLFRLSLAN
nchar(Seq1093)
#> [1] 1093
data(HydroScore)
LongSeqScore <- CharSequence2ScoreSequence(Seq1093, HydroScore)</pre>
table(LongSeqScore)
#> LongSeqScore
#> -5 -4 -3 -2 -1
                       0
                           2 3
                                       5
                                    4
#> 81 222 22 82 232 85 78 103 157
localScoreC(LongSeqScore)
#> $localScore
#> value begin
               end
#>
     65 956 1001
#>
#> $suboptimalSegmentScores
#>
     value begin end
#> 1
        40
              1
                   20
#> 2
        10
              71
                   73
#> 3
        23
                   99
              80
#> 4
         3
             114 114
#> 5
         3
             124 124
#> 6
             128 128
         4
#> 7
        23
             130 150
#> 8
        16
             181 195
#> 9
         2
             217 217
             224 224
#> 10
         4
#> 11
        33
             229 243
#> 12
         5
             294 296
#> 13
         4
             301 301
#> 14
         6
             304 307
#> 15
        36
             312 337
             379 379
#> 16
         4
#> 17
             384 384
         4
#> 18
         2
             387 387
#> 19
        14
             390 397
#> 20
             411 411
         4
#> 21
         3
             413 413
#> 22
        25
             416 486
#> 23
         9
             523 527
             533 537
#> 24
         9
#> 25
             548 548
         4
#> 26
         3
             554 554
#> 27
         3
             563 563
#> 28
         4
             566 566
#> 29
        10
             574 576
#> 30 16 588 601
```

```
#> 31 3
             633
                  633
#> 32
             637
                  637
         2
#> 33
        12
             644
                  646
#> 34
         6
             654
                  655
#> 35
             661
        12
                  668
#> 36
         7
             680
                  684
#> 37
         3
             691
                  691
#> 38
         8
             697
                  702
#> 39
         5
             711
                  712
#> 40
             715
                  715
         4
             725
#> 41
         6
                  727
#> 42
         3
             740
                  740
#> 43
         5
             742
                  744
         2
#> 44
             746
                  746
#> 45
             748 748
#> 46
         4
             754
                  754
#> 47
         3
             756
                  756
#> 48
         3
             766
                  766
#> 49
         5
             773
                  774
#> 50
        10
             778
                  780
#> 51
             794
         4
                  794
#> 52
         2
             799
                  799
#> 53
         3
             807 807
#> 54
         3
             811 811
         7
#> 55
             813
                  815
#> 56
             822
         3
                  822
             832
#> 57
         9
                  835
#> 58
         3
             844
                  844
         7
#> 59
             848
                  851
#> 60
         6
             864
                  868
         3
             879 879
#> 61
#> 62
         2
             883 883
#> 63
        16
             890 894
#> 64
        2
             924
                  924
#> 65
         3
             931 931
#> 66
             934 942
        13
#> 67
         3
             951 951
#> 68
         65
             956 1001
#> 69
         9 1048 1049
#> 70
         2 1054 1054
#> 71
         3 1056 1056
         4 1059 1059
#> 72
         4 1064 1064
#> 73
#> 74
         4 1074 1074
         3 1079 1079
#> 75
#> 76
         2 1083 1083
#> 77
         6 1089 1090
#>
#> $RecordTime
   [1]
               70
                    77
                         78
                              79
                                  112 113 115
                                                116 117 119
                                                               120
                                                                    121
                                                                         122
                                                                              123
#> [16]
                                       178
                                            179
                                                           219
                                                                220
                                                                     221
                                                                         222
                                                                              223
         125
              126
                   127
                        129
                             176
                                  177
                                                 180
                                                      218
#> [31]
         226
              227
                   228
                        290
                             291
                                  293
                                       299
                                            300
                                                 303
                                                      310
                                                           311
                                                               378
                                                                    381
                                                                         382
                                                                              383
         386 388 389
                                            531
                                                 532 550 551
#> [46]
                        412 415
                                  521
                                       522
                                                               558
                                                                    559
                                                                         560
                                                                              561
```

```
[61]
          562
                564
                     565
                           569
                                570
                                      571
                                           573
                                                 583
                                                      584
                                                            585
                                                                 586
                                                                       587
                                                                            631
                                                                                  632
                                                                                       635
    [76]
           636
                639
                      640
                           643
                                      652
                                           653
                                                 657
                                                      658
                                                                 660
                                                                       679
                                                                            687
                                                                                  688
                                                                                       689
                                651
                                                            659
    [91]
           690
                692
                      693
                           695
                                 696
                                      709
                                           710
                                                 714
                                                      716
                                                            717
                                                                 718
                                                                       719
                                                                            720
                                                                                  721
                                                                                       722
   [106]
           723
                724
                      735
                           736
                                 737
                                      738
                                           739
                                                 741
                                                      750
                                                            751
                                                                 752
                                                                       758
                                                                            759
                                                                                  760
                                                                                       761
   [121]
           762
                763
                      764
                           765
                                 769
                                      771
                                           777
                                                 791
                                                      792
                                                            793
                                                                 798
                                                                       803
                                                                            804
                                                                                  805
                                                                                       806
   [136]
           809
                810
                     812
                           819
                                820
                                      821
                                           824
                                                 825
                                                      826
                                                            827
                                                                 828
                                                                       829
                                                                            843
                                                                                  845
                                                                                       846
   [151]
           847
                863
                      874
                           875
                                876
                                           878
                                                 882
                                                      885
                                                            886
                                                                 888
                                                                       889
                                                                                       926
                                      877
                                                                            922
                                                                                  925
   [166]
           927
                928
                     929
                           930
                                932
                                      933
                                           949
                                                 950
                                                      954
                                                            955 1045 1046 1047 1052 1053
#> [181] 1055 1061 1062 1063 1069 1070 1071 1072 1073 1076 1077 1078 1080 1081 1082
   [196] 1085 1087 1088 1093
head(sort(localScoreC(LongSeqScore)$suboptimalSegmentScores[,1], decreasing = TRUE))
#> [1] 65 40 36 33 25 23
plot(1:1093,lindley(LongSeqScore), type = 'l')
```



plot(1:400,lindley(LongSeqScore)[1:400], type = 'l')



We get: local score = 65. Let us compute its p-value in a I.I.D. model.

```
LS <- localScoreC(LongSeqScore)$localScore["value"]
prob1 <- scoreSequences2probabilityVector(list(LongSeqScore))</pre>
prob1
#>
           -5
                                  -3
                                             -2
                                                         -1
                                                                     0
#> 0.07410796 0.20311070 0.02012809 0.07502287 0.21225984 0.07776761 0.00000000
            2
                       3
                                              5
#> 0.07136322 0.09423605 0.14364135 0.02836231
daudin(local_score = LS, sequence_length = length(LongSeqScore),
       score_probabilities = prob1,
       sequence_min = min(LongSeqScore),
       sequence_max = max(LongSeqScore))
#> [1] 0.07231933
```

In an independent model the p-value equals 7.2%, that is non significant using the nominal level 5%. Let us compute it in a markovian model. We need the transition matrix.

```
tmpMarkovParameters <- sequences2transmatrix(LongSeqScore)
lambda <- tmpMarkovParameters$transition_matrix
apply(lambda, 1, sum) #to check stochasticity
#> -5 -4 -3 -2 -1 0 2 3 4 5
#> 1 1 1 1 1 1 1 1 1 1
prob0 <- stationary_distribution(lambda)
print(prob0)
#> [1] 0.07415553 0.20331534 0.02016219 0.07512607 0.21241680 0.07789931
#> [7] 0.07052191 0.09425453 0.14375820 0.02839011
score_values <- tmpMarkovParameters$score_value
print(score_values)
#> [1] -5 -4 -3 -2 -1 0 2 3 4 5
```

Notice that the score value 1 is not present in the sequence.

```
exact_mc(LS, lambda, sequence_length = length(LongSeqScore), score_values = score_values)
#> [1] 0.07695456
```

The p-value in a Markovian model, equal to 7.7% is very similar in this case with the one of the independent model.

Sub optimal excursions

Let us consider a study on the first and the last sub optimal excursions.

```
subOptSegment <- localScoreC(LongSeqScore)$suboptimalSegmentScores</pre>
o <- order(subOptSegment[,1], decreasing = TRUE)</pre>
subOptSegment <- subOptSegment[o,] # reordering segments by decreasing score values
print(subOptSegment)
#>
      value begin end
#> 68
         65
              956 1001
#> 1
         40
                    20
                1
#> 15
         36
              312 337
#> 11
         33
              229 243
#> 22
         25
              416 486
#> 3
         23
               80
                   99
#> 7
         23
              130 150
#> 8
         16
              181 195
#> 30
         16
              588 601
#> 63
         16
              890 894
#> 19
         14
              390 397
#> 66
         13
              934 942
              644 646
#> 33
         12
#> 35
         12
              661
                  668
#> 2
              71
         10
                   73
#> 29
         10
              574 576
#> 50
         10
              778
                  780
#> 23
          9
              523 527
#> 24
          9
              533 537
#> 57
              832 835
          9
          9 1048 1049
#> 69
#> 38
              697 702
          8
#> 36
          7
              680 684
#> 55
          7
              813 815
#> 59
          7
              848 851
#> 14
          6
              304
                  307
#> 34
          6
              654 655
#> 41
          6
              725 727
#> 60
          6
              864
                  868
#> 77
            1089 1090
          6
#> 12
          5
              294 296
#> 39
          5
              711
                   712
#> 43
          5
              742 744
#> 49
              773 774
              128 128
#> 6
              224
                  224
#> 10
```

```
#> 13
                301
                     301
#> 16
                379
                     379
           4
#> 17
                384
                     384
#> 20
                411
                     411
#> 25
                548
                     548
#> 28
                566
                     566
#> 40
                715
                     715
#> 45
                748
                     748
                754
                     754
#> 46
                794
#> 51
           4
                     794
#> 72
               1059 1059
#> 73
               1064 1064
           4
#> 74
               1074 1074
           4
#> 4
                114
           3
                     114
           3
#> 5
                124
                     124
#> 21
           3
                413
                     413
#> 26
           3
                554
                     554
#> 27
           3
                563
                     563
#> 31
           3
                633
                     633
#> 37
           3
                691
                     691
#> 42
           3
                740
                     740
#> 47
           3
                756
                     756
           3
                     766
#> 48
                766
#> 53
           3
                807
                     807
           3
#> 54
                811
                     811
#> 56
           3
                822
                     822
#> 58
           3
                844
                     844
#> 61
           3
                879
                     879
#> 65
           3
                931
                     931
           3
                951
                     951
#> 67
#>
   71
           3
               1056 1056
#>
   75
           3
               1079 1079
#> 9
           2
                217
                     217
#> 18
           2
                387
                     387
#> 32
           2
                637
                     637
           2
#> 44
                746
                     746
#> 52
           2
                799
                     799
#> 62
           2
                883
                     883
#> 64
           2
                924
                     924
#> 70
           2
               1054 1054
#> 76
           2
               1083 1083
```

The sub optimal scores are 40, 36 and 33 realized by the "visual" excursions number 1, 15 and 11. Such number of excursion, 11 and 15 are large enough to avoid considering the mathematical number. The first excursion is mathematically the first one (see below).

```
lindley(LongSeqScore)
#>
      [1]
          2
            6 10
                  8
                     8
                        6 10 12 16 15 19 23 27 26 25 29 28 32 36 40 39 36 40 39
     [25] 38 37 33 29 34 36 35 31 30 29 28 24 20 16 18 21 20 15 10 6
#>
                                                                      1 6 10
     [49] 12 15 11 7 11 7
                                                                   3
#>
                           5
                              4 8 11 7 10 9
                                               8
                                                  6
                                                     6 10
                                                           5
                                                              1
                                                                4
                                                                      0
                                                                            6
                                               6
                                                  5
#>
     [73] 10 6
               4 3 0 0
                           0
                              4 9 13 13 15 10
                                                     9 12
                                                           7 11 10 14 16 12 16
#>
     [97] 15 19 23 19 21 20 16 15 17 16 15 11
                                            7 6
                                                  1
                                                     0
                                                        0
                                                           3 0 0
                                                        7 11 14 16 19 19 18 14
    [121]
               0 3 0 0 0 4 0 4 8 13 17 16 16 11
#>
   [145] 16 19 18 16 20 23 22 21 16 12 16 16 12 16 15 10 9 14 10 6 11 7 7 11
```

```
[169] 13 8 11 9 8 4 2 0 0 0 0 0 2 6 11 10 9 5 5 1 5 4 6 8
    [193] 7 11 16 12 15 14 14 9 5 3 5 10 9 4 3 7 7 6 6 4 2 6 1 0
    [217] 2 0 0 0 0 0 0 4 0 0 0 0 3 7 9 11 14 10 15 15 19 22 24 23
    [241] 26 29 33 28 24 20 22 26 22 19 15 18 18 13 12 16 15 14 9 13 15 10 14 17
#>
    [265] 13 9 5 9 9 9 4 7 11 15 11 7 6 5 4 3 6 8 4 6 1 5 1 4
    [289] 3 0 0 0 0 4 2 5 4 3 0 0 4 0 0 2 5 2 6 4 0 0 0 4
#>
    [313] 9 8 8 12 15 14 13 9 13 9 8 13 15 17 16 18 22 25 27 30 26 30 29 31
#>
    [337] 36 35 31 33 36 32 32 30 33 28 27 23 19 15 13 8 10 12 11 15 13 18 20 16
   [361] 14 19 17 13 16 12 15 15 14 18 16 12 11 11 9 5 1 0 4 3 0 0 0
#>
    [385] 0 0 2 0 0 4 7 11 13 12 8 10 14 10 8 12 11 9 5 3
                                                                  6 10 9 5
    [409] 1 0 4 0 3 2 0 4 8 12 8 12 16 12 14 10 6 5 9 8 5 9 13 12
   [433] 17 17 16 12 11 11 10 15 19 15 17 21 20 19 21 20 15 14 18 13 13 16 15 19
#>
    [457] 15 11 15 12 16 20 18 16 16 11 15 11 9 13 8 7 11 6 11 15 12 11 13 8
#>
    [481] 10 14 17 21 21 25 24 20 15 19 23 18 22 20 24 20 15 18 17 19 18 15 14 10
    [505] 10 12 15 19 19 21 16 12 10 9 12 12 11 7 7 3 0 0 4 7 6 5 9 5
#>
    [529] 1 0 0 0 2 1 5 4
                                9
                                   5
                                     1
                                         6
                                           5
                                              4
                                                 7 5 9 4
                                                            0
                                                               4
                                                                  3
                           0
                                                    0 0 0 0
    [553]
         0 3
               3 1 0
                        0
                             0
                                0
                                   0
                                      3
                                         0
                                           0
                                              4
                                                 0
                                                               0
                                                                     3
    [577] 5 7
               2
                        0
                           0
                             0
                                0
                                   0
                                         3
                                           3
                                              3
                                                 7
                                                    3 6 10 10
                                                                  7 12 9 14
#>
                 1
                                      0
                                                               8
                             7
    [601] 16 12 15 14
                     9
                        5
                           5
                                6
                                   5
                                      7
                                         6
                                           5
                                              4
                                                 3
                                                    5
                                                      1 4 3
                                                                  1 4 4
                     7
                           0
                                3
                                            2
                                              0
                                                       0 0 0 5 8 12 12
#>
    [625] 9 12
               8 12
                        3
                             0
                                   2
                                      0
                                         0
                                                 0
                                                    0
    [649]
          7
             2
                0
                  0
                     0
                        3
                           6
                             2
                                0
                                   0
                                      0
                                         0
                                            3
                                              1
                                                 5
                                                    3
                                                      8 11 10 12 11 10 10
#>
             7
#>
    [673]
          8
               3
                  6
                     5
                        1
                           0
                             3
                                3
                                   3
                                      3
                                         7
                                           3
                                              2
                                                 0
                                                    0
                                                      0 0 3
                                                               0
                                                                  0
                                                                     0
                             7
                                3
                                   2
                                      5
                                              0
                                                 2
                                                    5
                                                      3 0
#>
    [697]
          3
            1
                1
                  4
                     4
                        8
                           4
                                         1
                                            0
                                                            4
                                                                  0
    [721]
                0
                        2
                           6
                             2
                                6
                                              3
                                                 0
                                                    0
                                                      0
                                                         0
                                                            0
                                                               3
#>
          0
             0
                  0
                                   5
                                      1
                                         1
                                            1
                                                                  0
                                                                     3
                                                                        2
                     4
             2
                0
                        0
                           0
                             0
                                         3
                                           3
                                              0
                                                 0
                                                    0
                                                       0
                                                         0
                                                            0
#>
    [745]
          0
                  4
                     2
                                0
                                   4
                                      0
                                                               0
                                                                  0
                                                                     3
                                                                           3
    [769]
          0
             0
                0
                  0
                     2
                        5
                          1
                             0
                                0
                                      6 10
                                            6
                                              2
                                                 6 10
                                                      5
                                                               3
                                                                  2
#>
                                   2
#>
    [793]
          0
            4
                0
                  0
                     0
                        0
                          2
                             1
                                0
                                   0
                                      0
                                         0
                                            0
                                              0
                                                 3
                                                    2
                                                       0
                                                         0
                                                            3
                                                               0
   [817]
          6
                0
                  0
                     0
                        3
                           2
                             0
                                0
                                         0
                                            0
                                              0
                                                 0
                                                    4
                                                       2
                                                         5
                                                            9
                                                               9
                                                                  5
                                                                     7
#>
             1
                                   0
                                      0
                                                                        9
#>
    [841]
          4
             0
               0
                  3
                     0
                        0
                           0
                             2
                                5
                                   3
                                      7
                                         2
                                           2
                                              4
                                                 3
                                                    2 5
                                                            3
                                                               5
                                                                     3
                                                                        0
                                                         4
                                                                  4
                                                                           .3
               3 6 6
                             3
                                2
                                                3 2 1 0
                                                            2
                                                               0
#>
    [865]
          2 4
                       6
                          6
                                   0
                                      0
                                        0
                                           0
                                              0
#>
   [889]
         0 5 8 12 12 16 13 12
                                8
                                   4
                                      6 10 13 12 11
                                                    7 9 12 10
                                                               6
                                                                  6
                                                                     5
         7 6 2 1 1 5 8
                             7
                                3
   [913]
                                   0
                                      0
                                        2
                                           0
                                              0
                                                 0
                                                    0 0 0
                                                            3
                                                               0
   [937] 6 4 4 6 9 13 13
                             9
                                8 7 3 2 0 0 3
                                                   1 0 0 0 5 3 7 12 16
#>
   [961] 14 16 15 14 18 14 10 9 8 7 10 10 10 13 9 13 18 15 19 24 28 27 27 31
   [985] 30 33 36 40 44 43 43 47 51 50 54 56 60 59 63 62 65 61 58 61 57 52 48 47
#> [1009] 43 39 38 37 41 45 42 40 42 41 39 35 32 36 33 32 28 28 28 28 27 25 21 17
#> [1033] 13 9 8 7 5 7 3 6 2 1 5 1 0 0 0 4 9 7 3 0 0 2 0 3
#> [1057] 2 0 4 0 0 0 0 4 3 2 1 0 0 0 0 0 4 0 0 0
#> [1081] 0 0 2 1 0 0 0 0
                                3 6 4
                                        0 0
recordTimes(LongSeqScore)
                                113 115 116
    [1]
          70
              77
                   78
                        79
                           112
                                              117 119
                                                        120
                                                            121
                                                                 122
                                                                      123
                                                                           125
#>
   [16]
         126
             127
                  129
                       176
                           177
                                178
                                     179
                                          180
                                              218
                                                   219
                                                        220
                                                             221
                                                                 222
                                                                           226
                       291
#>
   [31]
         227
             228
                  290
                            293
                                299
                                     300
                                          303
                                              310
                                                   311
                                                        378
                                                             381
                                                                 382
                                                                      383
                                                                           386
#>
   [46]
         388
             389
                  412
                       415
                            521
                                 522
                                     531
                                          532
                                               550
                                                   551
                                                        558
                                                             559
                                                                 560
                                                                      561
                                                                           562
              565
                            571
                                 573
                                     583
                                          584
                                               585
                                                        587
#>
   [61]
         564
                  569
                       570
                                                   586
                                                             631
                                                                 632
                                                                      635
                                                                           636
#>
   [76]
         639
              640
                  643
                       651
                            652
                                653
                                     657
                                          658
                                               659
                                                   660
                                                        679
                                                             687
                                                                 688
                                                                      689
                                                                           690
   [91]
         692
              693
                  695
                       696
                            709
                                 710
                                     714
                                          716
                                               717
                                                   718
                                                        719
                                                             720
                                                                  721
                                                                      722
                                                                           723
#> [106]
         724
              735
                  736
                       737
                            738
                                 739
                                     741
                                          750
                                               751
                                                   752
                                                        758
                                                             759
                                                                 760
                                                                      761
                                                                           762
                                     791
                                               793
#> [121]
         763
             764
                  765
                       769
                            771
                                 777
                                          792
                                                   798
                                                        803
                                                             804
                                                                 805
                                                                      806
                                                                           809
#> [136]
             812
                            821
                                 824
                                          826
                                               827
                                                   828
                                                        829
        810
                  819
                       820
                                     825
                                                             843
                                                                 845
                                                                      846
                                                                           847
#> [151]
         863
             874
                  875
                       876
                            877
                                878
                                     882
                                          885
                                              886
                                                   888
                                                        889
                                                            922
                                                                           927
                                                                 925
                                949 950 954
#> [166] 928 929
                  930
                       932
                           933
                                              955 1045 1046 1047 1052 1053 1055
#> [181] 1061 1062 1063 1069 1070 1071 1072 1073 1076 1077 1078 1080 1081 1082 1085
```

```
#> [196] 1087 1088 1093
LongSeqScore[1] > 0
#> [1] TRUE
```

Let us consider a study on the first and the last sub optimal excursions in the sequential order of the sequence.

First excursion height equal to 40; last one, the 77th one, height equal to 6.

```
subOptSegment["1",] #First excursion
#> value begin end
#> 1     40     1     20
subOptSegment[as.character(dim(subOptSegment)[1]),] #Last excursion
#> value begin end
#> 77     6     1089     1090
```

Let us compute their p-values.

```
theta <- letters[1:length(score_values)] # arbitrary</pre>
score_function <- score_values</pre>
                                           # defined earlier
a <- 40
i <- 1
system.time(pv2<-proba_theoretical_ith_excursion_markov(a, theta, lambda,
                                         score_function,i)$proba_q_i_geq_a)
#> utilisateur
                    système
                                 écoulé
#>
        34.650
                      0.001
                                 34.661
pv2
#> [1] 0.004478564
a <- 6
i <- 77
system.time(pv3<-proba_theoretical_ith_excursion_markov(a, theta, lambda,
                                         score_function, i)$proba_q_i_geq_a)
#> utilisateur
                    système
                                 écoulé
         1.593
                                  1.606
#>
                      0.013
pv3
#> [1] 0.1741784
```

First excursion : height equal to 40 with p-value=0.45%; Last excursion, the 77th one, height equal to 6 and p-value=17%.

The time computation of the first p-value is larger because of the larger value of a.

We can consider that from the 20th mountain we reached the stationary distribution for the beginning of excursion. We shall therefore take a lower value for i for the last excursion to evaluate the difference. Let us try i = 20 instead of 77.

We obtain the same value as expected even for i = 10.

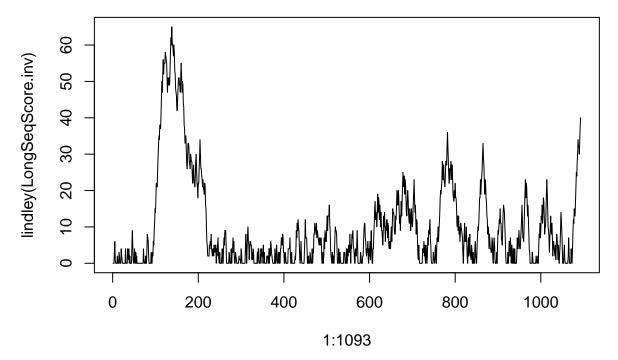
With a reverse lecture of the protein

As the lecture of a protein could be done in both direction, we also consider the reverse sequence.

```
LongSeqScore.inv <- rev(LongSeqScore)</pre>
localScoreC(LongSeqScore.inv)
#> $localScore
#> value begin
                end
#>
     65 93
               138
#>
#> $suboptimalSegmentScores
     value begin end
#> 1
        6
                   5
             4
#> 2
         2
              11
                  11
#> 3
         3
             15 15
#> 4
         4
             20 20
#> 5
             30 30
         4
#> 6
             35 35
         4
#> 7
         2
             40 40
#> 8
         9
             45 46
#> 9
         4
             51 51
#> 10
         3
             54
                 54
         2
#> 11
             56 56
#> 12
             72
                 72
         4
#> 13
         2
             77 77
#> 14
         8
              80 81
#> 15
         3
             90 90
#> 16
        65
             93 138
         7
#> 17
             243 246
#> 18
         3
             250 250
#> 19
         4
             255 256
#> 20
         9
             259 262
         3
#> 21
             272 272
#> 22
         4
             277 277
#> 23
            279 281
         7
#> 24
         3
             287 287
#> 25
             295 295
#> 26
             300 300
         4
#> 27 8
             310 311
```

```
#> 28
      10
             314 316
#> 29
             338
                  338
         3
#> 30
         4
             340 340
#> 31
             346 346
         4
#> 32
             350
                  352
         5
#> 33
         3
             354
                   354
#> 34
         2
             360
                  360
#> 35
             365
                  365
         4
#> 36
             367
                  369
         6
#> 37
             379
                  379
         4
#> 38
             382
         5
                  383
#> 39
         3
             387
                  387
#> 40
         3
             390
                  390
             392
#> 41
         8
                  397
#> 42
        3
             403 403
        7
#> 43
             410 414
#> 44
         3
             418 418
#> 45
        12
             426 433
#> 46
             439 440
        6
#> 47
        12
             448 450
#> 48
        2
             457 457
        3
#> 49
             461 461
#> 50
         4
             466 466
#> 51
             468 472
        11
             491
#> 52
         3
                  491
             493 506
#> 53
        16
#> 54
        3
             513 513
#> 55
        2
             516 516
#> 56
        10
             518
                  520
#> 57
             528
         4
                  528
#> 58
             531
         3
                  531
#> 59
         3
             540
                  540
#> 60
             546 546
         4
#> 61
         8
             549
                  554
#> 62
         9
             557
                  561
#> 63
         9
             567
                  571
#> 64
         3
             579
                  579
#> 65
        11
             584
                  588
#> 66
         9
             599
                  604
              608
#> 67
         25
                  678
#> 68
             724
                  727
         6
         5
#> 69
             732
                  732
#> 70
              735
                  741
        12
#> 71
         3
              749
                  749
#> 72
         5
             753
                  754
#> 73
        36
              757
                  782
#> 74
             820
                  822
        11
             830
#> 75
         8
                  834
#> 76
         3
             842
                  842
#> 77
         6
             846
                  847
#> 78
         33
             851
                   865
#> 79
         7
             890
                  891
#> 80
             897 897
         3
```

```
#> 81
         16
               899
                    913
#> 82
          3
               923
                    923
#> 83
          7
               925
                    929
#> 84
               932
                    932
          5
#> 85
                    936
          4
               936
#> 86
          4
               939
                    939
#> 87
         23
               944
                    964
#> 88
          3
               980
                    980
#> 89
          2
               989
                    989
#> 90
          2
               993
                    993
#> 91
         23
               995 1014
#> 92
         14
              1044 1047
#> 93
          5
              1054 1055
          7
#> 94
              1064 1065
#> 95
             1071 1071
          4
#> 96
         40
             1074 1093
#>
#> $RecordTime
                                        9
                                            10
#>
     [1]
             0
                  1
                        2
                             3
                                  7
                                                  12
                                                       13
                                                                  18
                                                                       19
                                                                             23
                                                                                  24
                                                                                        25
                                                             14
                                                       42
#>
    [16]
            26
                 27
                      28
                            29
                                 32
                                       33
                                            34
                                                 41
                                                                       49
                                                                             50
                                                                                  53
                                                                                        55
                                                             43
                                                                  44
    [31]
                                                                  71
                                                                                        79
#>
            58
                 59
                      60
                                            64
                                                  65
                                                       69
                                                             70
                                                                       74
                                                                             75
                                                                                  76
                            61
                                 62
                                       63
#>
    [46]
           85
                 86
                      87
                            88
                                 89
                                       92
                                           249
                                                 251
                                                      252
                                                           253
                                                                 254
                                                                      267
                                                                            268
                                                                                 269
                                                                                       270
#>
    [61]
          271
                274
                      275
                           276
                                278
                                      285
                                           286
                                                 291
                                                      293
                                                           294
                                                                 299
                                                                      303
                                                                            304
                                                                                 305
                                                                                       306
    [76]
#>
          308
                309
                     330
                           331
                                332
                                      333
                                           334
                                                 335
                                                      336
                                                           339
                                                                 343
                                                                      344
                                                                            345
                                                                                 349
                                                                                       355
    [91]
          356
                357
                     358
                           359
                                363
                                      364
                                           374
                                                 375
                                                      376
                                                                 378
                                                                                       389
#>
                                                           377
                                                                      381
                                                                            385
                                                                                 386
#> [106]
          391
                401
                     402
                           405
                                406
                                      407
                                           408
                                                 409
                                                      416
                                                           417
                                                                 419
                                                                      420
                                                                            421
                                                                                 422
                                                                                       424
#> [121]
          425
                438
                     442
                           443
                                444
                                      445
                                           446
                                                455
                                                      456
                                                           459
                                                                 460
                                                                      463
                                                                            464
                                                                                 465
                                                                                       489
#> [136]
          490
                492
                     512
                           515
                                517
                                      527
                                           530
                                                533
                                                      534
                                                           535
                                                                 536
                                                                      537
                                                                           538
                                                                                 543
                                                                                       544
   [151]
          545
                548
                     565
                           566
                                574
                                      576
                                           577
                                                 582
                                                      583
                                                           598
                                                                 607
                                                                      717
                                                                            718
                                                                                 719
                                                                                       721
#> [166]
                723
                     731
                           734
                                745
                                      746
                                           747
                                                 748
                                                      752
                                                           819
                                                                 829
                                                                      840
                                                                            843
                                                                                 844
                                                                                       845
          722
#> [181]
          849
                850
                     894
                           896
                                898
                                      920
                                           921
                                                 922
                                                      924
                                                           931
                                                                 934
                                                                      935
                                                                            941
                                                                                 942
                                                                                      943
#> [196] 975
               977 978
                          979
                                982
                                      983
                                           984
                                                985
                                                      986
                                                           987
                                                                 988
                                                                      991
                                                                            992
                                                                                 994 1043
#> [211] 1052 1053 1057 1058 1059 1060 1061 1062 1063 1067 1068 1069 1070
head(sort(localScoreC(LongSeqScore.inv)$suboptimalSegmentScores[,1], decreasing = TRUE))
#> [1] 65 40 36 33 25 23
plot(1:1093,lindley(LongSeqScore.inv), type = 'l')
```



That leads to: first excursion equal to 6; last excursion, the 96th one, equal to 38.

The corresponding p-values are:

```
markov_parameters <- sequences2transmatrix(LongSeqScore.inv)</pre>
lambda.inv <- markov_parameters$transition_matrix</pre>
system.time(pv5<-proba_theoretical_ith_excursion_markov(a = 38, theta,
                                         lambda.inv,
                                         score_function, i = 96
                                         ) $proba_q_i_geq_a)
#> utilisateur
                    système
                                  écoulé
#>
        31.748
                      0.004
                                  31.760
pv5
#> [1] 0.005460216
proba_theoretical_ith_excursion_markov(a = 6, theta, lambda.inv,
                                         score_function,i = 1
                                         )$proba_q_i_geq_a
#> [1] 0.1765172
```

Last excursion, the 96th one, equal to 38, with p-value equal to 0.53%. First excursion equal to 6, with p-value equal to 17%.

The last excursion is still significant. The first one is still non significant.

Remark: Even with a Bonferroni correction to take into account the multiple test (here two studies excursions), the sequence possesses a significant segment. Whereas considering the highest value over all the excursions of the whole sequence, the local score value, the sequence is not significant.

What about the excursion realising the local score

Let us consider the excursion 30 as an excursion among the others. For the first way to read the protein:

There is a less than 4 in 10,000 chance of having a first mountain over 65.

Or with the second way to read the protein:

As expected we obtain a similar p-value.

Although the highest mountain of height 65 in this sequence of length 1093 does not have a significant value at the 5% threshold, it does not mean that the sequence has not interesting segments. Observing a mountain exceeding 40 is significant for example, and it is the same for values 36 and 33. All these tests remain significant even if a correction of type Bonferroni is taken.