Follow-up data with the Epi package

Spring 2009.

Michael Hills Retired

Highgate, London

Martyn Plummer International Agency for Research on Cancer, Lyon

plummer@iarc.fr

Bendix Carstensen Steno Diabetes Center, Gentofte, Denmark

& Department of Biostatistics, University of Copenhagen

bxc@steno.dk

www.pubhealth.ku.dk/~bxc

Contents

1	Follow-up data in the Epi package	1
2	Timescales	1
3	Splitting the follow-up time along a timescale	4
4	Splitting time at a specific date	6
5	Competing risks — multiple types of events	8
6	Multiple events of the same type (recurrent events)	10

```
> library(Epi)
> print( sessionInfo(), l=F )
R version 3.1.0 (2014-04-10)
Platform: i386-w64-mingw32/i386 (32-bit)

attached base packages:
[1] utils datasets graphics grDevices stats methods base
other attached packages:
[1] Epi_1.1.64 foreign_0.8-61

loaded via a namespace (and not attached):
[1] tools_3.1.0
```

1 Follow-up data in the Epi package

In the Epi-package, follow-up data is represented by adding some extra variables to a dataframe. Such a dataframe is called a Lexis object. The tools for handling follow-up data then use the structure of this for special plots, tabulations etc.

Follow-up data basically consists of a time of entry, a time of exit and an indication of the status at exit (normally either "alive" or "dead"). Implicitly is also assumed a status *during* the follow-up (usually "alive").

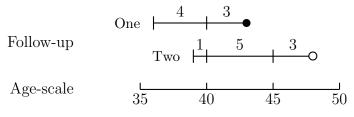


Figure 1: Follow-up of two persons

2 Timescales

A timescale is a variable that varies deterministically within each person during follow-up, e.g.:

- Age
- Calendar time
- Time since treatment
- Time since relapse

All timescales advance at the same pace, so the time followed is the same on all timescales. Therefore, it suffices to use only the entry point on each of the time scale, for example:

2 Timescales

- Age at entry.
- Date of entry.
- Time since treatment (at treatment this is 0).
- Time since relapse (at relapse this is 0)...

In the Epi package, follow-up in a cohort is represented in a Lexis object. A Lexis object is a dataframe with a bit of extra structure representing the follow-up. For the nickel data we would construct a Lexis object by:

The entry argument is a named list with the entry points on each of the timescales we want to use. It defines the names of the timescales and the entry points. The exit argument gives the exit time on one of the timescales, so the name of the element in this list must match one of the neames of the entry list. This is sufficient, because the follow-up time on all time scales is the same, in this case ageout - agein. Now take a look at the result:

```
> str( nickel )
  'data.frame':
                        679 obs. of 7 variables:
   $ id
            : num 3 4 6 8 9 10 15 16 17 18 ...
   $ icd
             : num 0 162 163 527 150 163 334 160 420 12 ...
   $ exposure: num 5 5 10 9 0 2 0 0.5 0 0 ...
             : num
                     1889 1886 1881 1886 1880 ...
                     17.5 23.2 25.2 24.7 30 ...
             : num
             : num 45.2 48.3 53 47.9 54.7 ...
   $ agein
   $ ageout : num 93 63.3 54.2 69.7 76.8 ...
> str( nicL )
  Classes 'Lexis' and 'data.frame':
                                             679 obs. of 14 variables:
             : num 1934 1934 1934 1934 ...
                     45.2 48.3 53 47.9 54.7
   $ age
             : num
                    27.7 25.1 27.7 23.2 24.8 ...
   $ tfh
             : num
   $ lex.dur : num
                    47.75 15 1.17 21.77 22.1 ...
   $ lex.Cst : num 0 0 0 0 0 0 0 0 0 0 ...
                     0 1 1 0 0 1 0 0 0 0 ...
   $ lex.Xst : num
   $ lex.id : int
                     1 2 3 4 5 6 7 8 9 10 ...
   $ id
             : num 3 4 6 8 9 10 15 16 17 18 ...
             : num 0 162 163 527 150 163 334 160 420 12 ...
   $ exposure: num 5 5 10 9 0 2 0 0.5 0 0 ...
             : num 1889 1886 1881 1886 1880 ...
                     17.5 23.2 25.2 24.7 30 ...
             : num
             : num 45.2 48.3 53 47.9 54.7 ...
   $ agein
   $ ageout : num 93 63.3 54.2 69.7 76.8 ...
   - attr(*, "time.scales")= chr "per" "age" "tfh"
- attr(*, "time.since")= chr "" "" ""
- attr(*, "breaks")=List of 3
    ..$ per: NULL
    ..$ age: NULL
    ..$ tfh: NULL
```

```
> head( nicL )
         per
                         tfh lex.dur lex.Cst lex.Xst lex.id id icd exposure
                 age
  1 1934.246 45.2273 27.7465 47.7535
                                                          1 3 0
                                           0
                                                   0
  2 1934.246 48.2684 25.0820 15.0028
                                           0
                                                   1
                                                          2 4 162
                                                                          5
  3 1934.246 52.9917 27.7465
                                           0
                                                          3 6 163
                                                                          10
                                           0
  4 1934.246 47.9067 23.1861 21.7727
                                                   0
                                                          4 8 527
                                                                          9
  5 1934.246 54.7465 24.7890 22.0977
                                           0
                                                   0
                                                          5 9 150
                                                                          0
  6 1934.246 44.3314 23.0437 18.2099
                                           0
                                                   1
                                                          6 10 163
                                                                          2
         dob
             age1st
                       agein
                             ageout
  1 1889.019 17.4808 45.2273 92.9808
  2 1885.978 23.1864 48.2684 63.2712
  3 1881.255 25.2452 52.9917 54.1644
  4 1886.340 24.7206 47.9067 69.6794
  5 1879.500 29.9575 54.7465 76.8442
  6 1889.915 21.2877 44.3314 62.5413
```

The Lexis object nicL has a variable for each timescale which is the entry point on this timescale. The follow-up time is in the variable lex.dur (duration).

There is a summary function for Lexis objects that list the numer of transitions and records as well as the total follow-up time:

```
> summary( nicL )
Transitions:
          To
From 0 1 Records: Events: Risk time: Persons:
          0 542 137 679 137 15348.06 679
```

We defined the exit status to be death from lung cancer (ICD7 162,163), i.e. this variable is 1 if follow-up ended with a death from this cause. If follow-up ended alive or by death from another cause, the exit status is coded 0, i.e. as a censoring.

Note that the exit status is in the variable lex.Xst (eXit status. The variable lex.Cst is the state where the follow-up takes place (Current status), in this case 0 (alive).

It is possible to get a visualization of the follow-up along the timescales chosen by using the plot method for Lexis objects. nicL is an object of class Lexis, so using the function plot() on it means that ${\bf R}$ will look for the function plot.Lexis and use this function.

```
> plot( nicL )
```

The function allows a lot of control over the output, and a points.Lexis function allows plotting of the endpoints of follow-up:

The results of these two plotting commands are in figure 2.

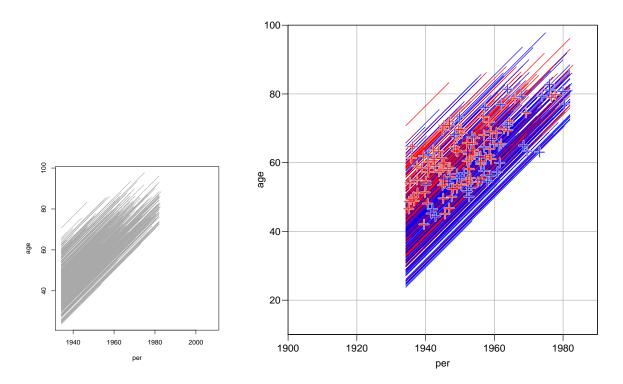


Figure 2: Lexis diagram of the **nickel** dataset, left panel the default version, the right one with bells and whistles. The red lines are for persons with exposure> 0, so it is pretty evident that the oldest ones are the exposed part of the cohort.

3 Splitting the follow-up time along a timescale

The follow-up time in a cohort can be subdivided by for example current age. This is achieved by the splitLexis (note that it is *not* called split.Lexis). This requires that the timescale and the breakpoints on this timescale are supplied. Try:

```
> nicS1 <- splitLexis( nicL, "age", breaks=seq(0,100,10) )</pre>
> summary( nicL )
  Transitions:
              1 Records:
                            Events: Risk time:
                                                 Persons:
     0 542 137
                       679
                                137
                                       15348.06
                                                       679
> summary( nicS1 )
  Transitions:
                  Records:
                             Events: Risk time:
                                                  Persons:
     0 2073 137
                       2210
                                        15348.06
                                                        679
                                 137
```

So we see that the number of events and the amount of follow-up is the same in the two datasets; only the number of records differ.

To see how records are split for each individual, it is useful to list the results for a few individuals:

```
12
        4 1936.34 50.00 25.28
                                  10.00
                                              0
                                                       0 8 527
                                                                       9 1886.34
        4 1946.34 60.00 35.28
                                  9.68
                                              0
                                                       0 8 527
                                                                       9 1886.34
13
14
        5 1934.25 54.75 24.79
                                  5.25
                                              0
                                                       0
                                                         9 150
                                                                       0 1879.50
15
        5 1939.50 60.00 30.04
                                  10.00
                                              0
                                                       0
                                                         9 150
                                                                       0 1879.50
        5 1949.50 70.00 40.04
                                              0
                                                       0 9 150
                                                                       0 1879.50
16
                                  6.84
17
        6 1934.25 44.33 23.04
                                  5.67
                                              0
                                                       0 10 163
                                                                       2 1889.91
        6 1939.91 50.00 28.71
                                              0
                                                                       2 1889.91
18
                                  10.00
                                                       0 10 163
19
        6 1949.91 60.00 38.71
                                              0
                                                       1 10 163
                                                                       2 1889.91
                                  2.54
   age1st agein ageout
    24.72 47.91
11
                 69.68
    24.72 47.91
12
                 69.68
13
    24.72 47.91
                 69.68
    29.96 54.75
                 76.84
14
15
    29.96 54.75
                  76.84
    29.96 54.75
16
                 76.84
    21.29 44.33
                 62.54
17
18
    21.29 44.33
                 62.54
19
    21.29 44.33
                 62.54
```

The resulting object, nicS1, is again a Lexis object, and so follow-up may be split further along another timescale. Try this and list the results for individuals 8, 9 and 10 again:

```
> nicS2 <- splitLexis( nicS1, "tfh", breaks=c(0,1,5,10,20,30,100) )</pre>
> round( subset( nicS2, id %in% 8:10 ), 2 )
     lex.id
                      age
                             tfh lex.dur lex.Cst lex.Xst id icd exposure
                                                                                dob
                per
          4 1934.25 47.91 23.19
                                                                          9 1886.34
  13
                                                           8 527
                                     2.09
                                                0
                                                         0
                                                0
                                                            8 527
                                                                          9 1886.34
  14
          4 1936.34 50.00 25.28
                                     4.72
                                                         0
  15
          4 1941.06 54.72 30.00
                                     5.28
                                                0
                                                         0
                                                            8 527
                                                                          9 1886.34
  16
          4 1946.34 60.00 35.28
                                     9.68
                                                0
                                                         0
                                                           8 527
                                                                          9 1886.34
  17
          5 1934.25 54.75 24.79
                                     5.21
                                                0
                                                         0
                                                           9 150
                                                                          0 1879.50
                                                0
          5 1939.46 59.96 30.00
                                                         Ω
                                                            9 150
                                                                          0 1879.50
  18
                                     0.04
  19
          5 1939.50 60.00 30.04
                                    10.00
                                                0
                                                         0
                                                            9 150
                                                                          0 1879.50
  20
          5 1949.50 70.00 40.04
                                     6.84
                                                0
                                                         0 9 150
                                                                          0 1879.50
          6 1934.25 44.33 23.04
  21
                                     5.67
                                                0
                                                         0 10 163
                                                                          2 1889 91
  22
          6 1939.91 50.00 28.71
                                                0
                                                                          2 1889.91
                                     1.29
                                                         0 10 163
  23
          6 1941.20 51.29 30.00
                                     8.71
                                                0
                                                         0 10 163
                                                                          2 1889.91
  24
          6 1949.91 60.00 38.71
                                     2.54
                                                         1 10 163
                                                                          2 1889.91
     age1st agein ageout
     24.72 47.91
  13
                   69.68
      24.72 47.91
  14
  15
      24.72 47.91
                   69.68
  16
      24.72 47.91
                    69.68
  17
      29.96 54.75
                    76.84
      29.96 54.75
  18
                    76.84
  19
      29.96 54.75
                    76.84
  20
      29.96 54.75
                    76.84
      21.29 44.33
                    62.54
  21
      21.29 44.33
  22
                    62.54
  23
      21.29 44.33
                    62.54
      21.29 44.33
                   62.54
```

If we want to model the effect of these timescales we will for each interval use either the value of the left endpoint in each interval or the middle. There is a function timeBand which returns these. Try:

```
id lex.id
                                    tfh lex.dur mid.age mid.tfh
                   per
                            age
           1 1934.246 45.2273 27.7465
1
                                          2.2535
                                                       45
    3
           1 1936.500 47.4808 30.0000
                                          2.5192
                                                       45
                                                                65
    3
              1939.019 50.0000 32.5192
                                        10.0000
                                                       55
                                                                65
           1 1949.019 60.0000 42.5192 10.0000
    3
                                                       65
                                                                65
           1 1959.019 70.0000 52.5192
                                         10.0000
                                                       75
                                                                65
6
    3
                                                       85
           1 1969.019 80.0000 62.5192
                                         10.0000
                                                                65
7
    3
           1 1979.019 90.0000 72.5192
                                          2.9808
                                                       95
                                                                65
8
    4
             1934.246 48.2684
                                25.0820
                                          1.7316
                                                       45
                                                                25
9
    4
           2 1935.978 50.0000 26.8136
                                                       55
                                                                25
                                          3.1864
10
    4
           2 1939.164 53.1864 30.0000
                                          6.8136
                                                       55
                                                                65
11
    4
           2 1945.978 60.0000 36.8136
                                          3.2712
                                                       65
                                                                65
12
    6
           3 1934.246 52.9917 27.7465
                                          1.1727
                                                       55
                                                                25
13
    8
            4 1934.246 47.9067
                                23.1861
                                          2.0933
                                                       45
                                                                25
14
                                                       55
                                                                25
    8
           4 1936.340 50.0000 25.2794
                                          4.7206
15
    8
           4 1941.060 54.7206 30.0000
                                          5.2794
                                                       55
                                                                65
16
    8
           4 1946.340 60.0000 35.2794
                                          9.6794
                                                       65
                                                                65
17
    9
           5 1934.246 54.7465 24.7890
                                          5.2110
                                                       55
                                                                25
    9
18
             1939.457 59.9575 30.0000
                                          0.0425
                                                       55
                                                                65
19
    9
           5 1939.500 60.0000 30.0425
                                         10.0000
                                                       65
                                                                65
20
           5 1949.500 70.0000 40.0425
                                          6.8442
```

Note that these are the midpoints of the intervals defined by breaks=, not the midpoints of the actual follow-up intervals. This is because the variable to be used in modelling must be independent of the consoring and mortality pattern — it should only depend on the chosen grouping of the timescale.

4 Splitting time at a specific date

If we have a recording of the date of a specific event as for example recovery or relapse, we may classify follow-up time as being before of after this intermediate event. This is achieved with the function cutlexis, which takes three arguments: the time point, the timescale, and the value of the (new) state following the date.

Now we define the age for the nickel vorkers where the cumulative exposure exceeds 50 exposure years:

```
> subset( nicL, id %in% 8:10 )
                          tfh lex.dur lex.Cst lex.Xst lex.id id icd exposure
                  age
  4 1934.246 47.9067 23.1861 21.7727
                                                     0
  5 1934.246 54.7465 24.7890 22.0977
                                             0
                                                     0
                                                                             0
                                                            5
                                                               9 150
  6 1934.246 44.3314 23.0437 18.2099
                                                            6 10 163
                                                                             2
             age1st
                        agein
                               ageout
  4 1886.340 24.7206 47.9067 69.6794
    1879.500 29.9575 54.7465 76.8442
  6 1889.915 21.2877 44.3314 62.5413
> agehi <- nicL$age1st + 50 / nicL$exposure</pre>
 nicC <- cutLexis( data=nicL, cut=agehi, timescale="age",</pre>
                    new.state=2, precursor.states=0 )
> subset( nicC, id %in% 8:10 )
                     age
                             tfh lex.dur lex.Cst lex.Xst lex.id id icd exposure
  4100 1934.246 47.9067 23.1861 21.7727
                                                                  8 527
                                                2
                                                        2
       1934.246 54.7465 24.7890 22.0977
                                                0
                                                        0
                                                                  9 150
                                                                                0
                                                0
                                                        2
       1934.246 44.3314 23.0437
                                  1.9563
                                                                6 10 163
                                                                                2
  680
       1936.203 46.2877 25.0000 16.2536
                                                                6 10 163
                 age1st
                           agein
                                  ageout
  4100 1886.340 24.7206 47.9067 69.6794
       1879.500 29.9575 54.7465 76.8442
```

```
6 1889.915 21.2877 44.3314 62.5413
680 1889.915 21.2877 44.3314 62.5413
```

(The precursor.states= argument is explained below). Note that individual 6 has had his follow-up split at age 25 where 50 exposure-years were attained. This could also have been achieved in the split dataset nicS2 instead of nicL, try:

```
> subset( nicS2, id %in% 8:10 )
     lex.id
                                 tfh lex.dur lex.Cst lex.Xst id icd exposure
                 per
                         age
  13
          4 1934.246 47.9067 23.1861
                                      2.0933
                                                    0
                                                              8 527
  14
          4 1936.340 50.0000 25.2794
                                      4.7206
                                                    0
                                                            0 8 527
  15
          4 1941.060 54.7206 30.0000
                                      5.2794
                                                    0
                                                            0 8 527
  16
          4 1946.340 60.0000 35.2794
                                      9.6794
                                                    0
                                                            0 8 527
                                                                            9
          5 1934.246 54.7465 24.7890
  17
                                      5.2110
                                                    0
                                                            0 9 150
                                                                            0
                                                            0 9 150
          5 1939.457 59.9575 30.0000
  18
                                      0.0425
                                                    0
                                                            0 9 150
                                                                            0
  19
          5 1939.500 60.0000 30.0425 10.0000
  20
          5 1949.500 70.0000 40.0425
                                      6.8442
                                                    0
                                                            0 9 150
                                                                            0
                                                                            2
  21
          6 1934.246 44.3314 23.0437
                                      5.6686
                                                    0
                                                            0 10 163
                                                                            2
  22
          6 1939.915 50.0000 28.7123
                                      1.2877
                                                    0
                                                            0 10 163
  23
          6 1941.203 51.2877 30.0000
                                      8.7123
                                                    0
                                                            0 10 163
  24
          6 1949.915 60.0000 38.7123
                                      2.5413
                                                    0
                                                            1 10 163
          dob age1st
                        agein ageout
  13 1886.340 24.7206 47.9067 69.6794
  14 1886.340 24.7206 47.9067 69.6794
  15 1886.340 24.7206 47.9067 69.6794
  16 1886.340 24.7206 47.9067 69.6794
  17 1879.500 29.9575 54.7465 76.8442
  18 1879.500 29.9575 54.7465 76.8442
  19 1879.500 29.9575 54.7465 76.8442
  20 1879.500 29.9575 54.7465 76.8442
  21 1889.915 21.2877 44.3314 62.5413
  22 1889.915 21.2877 44.3314 62.5413
  23 1889.915 21.2877 44.3314 62.5413
  24 1889.915 21.2877 44.3314 62.5413
> agehi <- nicS2$age1st + 50 / nicS2$exposure</pre>
> nicS2C <- cutLexis( data=nicS2, cut=agehi, timescale="age",
                      new.state=2, precursor.states=0 )
> subset( nicS2C, id %in% 8:10 )
       lex.id
                   per
                                    tfh lex.dur lex.Cst lex.Xst id icd exposure
                           age
  3142
            4 1934.246 47.9067 23.1861 2.0933
                                                              2 8 527
                                                      2
            4 1936.340 50.0000 25.2794
  3143
                                         4.7206
                                                                8 527
  3144
                                                      2
                                                              2 8 527
            4 1941.060 54.7206 30.0000
                                        5.2794
                                                                              9
  3145
            4 1946.340 60.0000 35.2794
                                         9.6794
                                                      2
                                                              2
                                                                 8 527
                                                                              9
  17
            5 1934.246 54.7465 24.7890
                                         5.2110
                                                      0
                                                              0
                                                                 9 150
                                                                              0
            5 1939.457 59.9575 30.0000
                                                      0
                                                              0
                                                                9 150
  18
                                         0.0425
                                                                              0
            5 1939.500 60.0000 30.0425 10.0000
  19
                                                                9 150
                                                      0
  20
            5 1949.500 70.0000 40.0425
                                         6.8442
                                                              0
                                                                9 150
                                                                              0
                                         1.9563
  21
            6 1934.246 44.3314 23.0437
                                                      0
                                                              2 10 163
                                                                              2
                                                              2 10 163
                                                                               2
  3150
            6 1936.203 46.2877 25.0000
                                         3.7123
            6 1939.915 50.0000 28.7123
                                                      2
                                                              2 10 163
                                                                              2
  3151
                                         1.2877
  3152
            6 1941.203 51.2877 30.0000
                                                      2
                                                              2 10 163
                                                                              2
                                         8.7123
  3153
            6 1949.915 60.0000 38.7123
                                         2.5413
                                                              1 10 163
            dob age1st
                          agein ageout
  3142 1886.340 24.7206 47.9067 69.6794
  3143 1886.340 24.7206 47.9067 69.6794
  3144 1886.340 24.7206 47.9067 69.6794
  3145 1886.340 24.7206 47.9067 69.6794
       1879.500 29.9575 54.7465 76.8442
  17
  18
       1879.500 29.9575 54.7465 76.8442
  19
       1879.500 29.9575 54.7465 76.8442
  20
       1879.500 29.9575 54.7465 76.8442
       1889.915 21.2877 44.3314 62.5413
```

```
3150 1889.915 21.2877 44.3314 62.5413 3151 1889.915 21.2877 44.3314 62.5413 3152 1889.915 21.2877 44.3314 62.5413 3153 1889.915 21.2877 44.3314 62.5413
```

Note that follow-up subsequent to the event is classified as being in state 2, but that the final transition to state 1 (death from lung cancer) is preserved. This is the point of the precursor.states= argument. It names the states (in this case 0, "Alive") that will be over-witten by new.state (in this case state 2, "High exposure"). Clearly, state 1 ("Dead") should not be updated even if it is after the time where the persons moves to state 2. In other words, only state 0 is a precursor to state 2, state 1 is always subsequent to state 2.

Note if the intermediate event is to be used as a time-dependent variable in a Cox-model, then lex.Cst should be used as the time-dependent variable, and lex.Xst==1 as the event.

5 Competing risks — multiple types of events

If we want to consider death from lung cancer and death from other causes as separate events we can code these as for example 1 and 2.

```
> data( nickel )
 nicL <- Lexis( entry = list( per=agein+dob,</pre>
                               age=agein,
                               tfh=agein-age1st ),
                  exit = list( age=ageout ),
           exit.status = (icd > 0) + (icd %in% c(162,163)),
                  data = nickel )
 summary( nicL )
  Transitions:
      To
  From 0
           1 2 Records:
                             Events: Risk time:
     0 47 495 137
                        679
                                 632
                                      15348.06
> subset( nicL, id %in% 8:10 )
                         tfh lex.dur lex.Cst lex.Xst lex.id id icd exposure
                 age
  4 1934.246 47.9067 23.1861 21.7727
                                           0
                                                 1
  5 1934.246 54.7465 24.7890 22.0977
                                           0
                                                          5
                                                             9 150
                                                                          0
                                                   1
  6 1934.246 44.3314 23.0437 18.2099
                                           0
                                                   2
                                                          6 10 163
                                                                          2
         dob age1st
                      agein ageout
  4 1886.340 24.7206 47.9067 69.6794
  5 1879.500 29.9575 54.7465 76.8442
  6 1889.915 21.2877 44.3314 62.5413
```

If we want to label the states, we can enter the names of these in the states parameter, try for example:

```
Transitions:
To
From Alive D.oth D.lung Records: Events: Risk time: Persons:
Alive 47 495 137 679 632 15348.06 679
```

Note that the Lexis function automatically assumes that all persons enter in the first level (given in the states= argument)

When we cut at a date as in this case, the date where cumulative exposure exceeds 50 exposure-years, we get the follow-up *after* the date classified as being in the new state if the exit (lex.Xst) was to a state we defined as one of the precursor.states:

```
> nicL$agehi <- nicL$age1st + 50 / nicL$exposure
 nicC <- cutLexis( data = nicL,
                     cut = nicL$agehi,
               timescale = "age",
               new.state = "HiExp"
       precursor.states = "Alive" )
 subset( nicC, id %in% 8:10 )
                             tfh lex.dur lex.Cst lex.Xst lex.id id icd exposure
            per
                    age
  4100 1934.246 47.9067 23.1861 21.7727
                                           HiExp
                                                   D.oth
       1934.246 54.7465 24.7890 22.0977
                                           Alive
                                                   D.oth
                                                               5 9 150
                                                                               0
       1934.246 44.3314 23.0437
                                  1.9563
                                           Alive
                                                   HiExp
                                                               6 10 163
                                                                               2
                                           HiExp
                                                  D.lung
  680
      1936.203 46.2877 25.0000 16.2536
                                                               6 10 163
                                                                                2
                          agein ageout
                                            agehi
            dob
                age1st
  4100 1886.340 24.7206 47.9067 69.6794 30.27616
       1879.500 29.9575 54.7465 76.8442
       1889.915 21.2877 44.3314 62.5413 46.28770
  680 1889.915 21.2877 44.3314 62.5413 46.28770
 summary( nicC, scale=1000 )
  Transitions:
       То
                                    Records:
                                               Events: Risk time:
          Alive HiExp D.oth D.lung
  From
                                                                    Persons:
                                 65
    Alive
             39
                   83
                        279
                                          466
                                                   427
                                                             10.77
                                                                         466
                                                                         296
              0
                    8
                                 72
                                          296
                                                   288
                                                              4.58
    HiExp
                        216
                         495
                                137
                                          762
                                                   715
                                                             15.35
                                                                         679
    Sum
```

Note that the persons-years is the same, but that the number of events has changed. This is because events are now defined as any transition from alive, including the transitions to HiExp.

Also note that (so far) it is necessary to specify the variable with the cutpoints in full, using only cut=agehi would give an error.

Subdivision of existing states

It may be of interest to subdivide the states following the intermediate event according to wheter the event has occurred or not. That is done by the argument split.states=TRUE.

Moreover, it will also often be of interest to introduce a new timescale indicating the time since intermediate event. This can be done by the argument new.scale=TRUE, alternatively new.scale="tfevent", as illustrated here:

```
split.states=TRUE, new.scale=TRUE,
       precursor.states = "Alive" )
> subset( nicC, id %in% 8:10 )
            per
                            tfh
                                   Hi.dur lex.dur lex.Cst
                                                              lex.Xst lex.id id icd
                    age
  4100 1934.246 47.9067 23.1861 17.63054 21.7727
                                                        Hi
                                                            D.oth(Hi)
                                                                            4
       1934.246 54.7465 24.7890
                                       NA 22.0977
                                                     Alive
                                                                D.oth
                                                                            5
                                                                              9 150
  6
       1934.246 44.3314 23.0437
                                       NA
                                           1.9563
                                                     Alive
                                                                   Ηi
                                                                            6 10 163
      1936.203 46.2877 25.0000
                                 0.00000 16.2536
                                                        Hi D.lung(Hi)
  680
                                                                            6 10 163
       exposure
                     dob age1st
                                    agein
                                           ageout
                                                      agehi
  4100
              9 1886.340 24.7206 47.9067 69.6794
                                                  30.27616
  5
              0 1879.500 29.9575 54.7465 76.8442
  6
                1889.915 21.2877 44.3314 62.5413 46.28770
              2 1889.915 21.2877 44.3314 62.5413 46.28770
> summary( nicC, scale=1000 )
  Transitions:
       То
          Alive Hi D.oth D.lung D.lung(Hi) D.oth(Hi)
  From
                                                        Records:
                                                                  Events:
                      279
    Alive
             39 83
                              65
                                          0
                                                     0
                                                             466
                                                                       427
    Ηi
              0 8
                       0
                              0
                                         72
                                                   216
                                                             296
                                                                       288
                      279
    Sum
             39 91
                              65
                                         72
                                                   216
                                                             762
                                                                       715
  Transitions:
          Risk time:
  From
                      Persons:
    Alive
               10.77
    Ηi
                4.58
                            296
    Sum
               15.35
                            679
```

6 Multiple events of the same type (recurrent events)

Sometimes more events of the same type are recorded for each person and one would then like to count these and put follow-up time in states accordingly. Essentially, each set of cutpoints represents progressions from one state to the next. Therefore the states should be numbered, and the numbering of states subsequently occupied be increased accordingly.

This is a behaviour different from the one outlined above, and it is achieved by the argument count=TRUE to cutLexis. When count is set to TRUE, the value of the arguments new.state and precursor.states are ignored. Actually, when using the argument count=TRUE, the function countLexis is called, so an alternative is to use this directly.