Follow-up data with the Epi package

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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	Foll	v-up data in the Epi package					
	1.1	Timescales	1				
	1.2	Splitting the follow-up time along a timescale	E				
	1.3	Splitting time at a specific date	7				
	1.4	Competing risks — multiple types of events	ę				
	1.5	Multiple events of the same type (recurrent events)	11				

Chapter 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").

These three variables are specific for each type of outcome, i.e. cancer, cardiovascular event, death, ...

1.1 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:

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

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:

2 1.1 Timescales

```
tfh=agein-age1st ),

exit = list( age=ageout ),

exit.status = ( icd %in% c(162,163) )*1,

data = nickel )
```

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. This is sufficient, because the follow-up time on all time scale is the same, in this case ageout - agein. Now take a look at the result:

```
scale is the same, in this case ageout - agein. Now take a look at the result:
> str( nickel )
'data.frame':
                     679 obs. of 7 variables:
         : 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 ...
 $ age1st : num 17.5 23.2 25.2 24.7 30.0 ...
         : num 45.2 48.3 53.0 47.9 54.7 ...
 $ ageout : num 93.0 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 ...
           : num 45.2 48.3 53.0 47.9 54.7 ...
 $ age
           : num 27.7 25.1 27.7 23.2 24.8 ...
$ tfh
 $ lex.dur : num 47.75 15.00 1.17 21.77 22.10 ...
 $ lex.Cst : num 0 0 0 0 0 0 0 0 0 ...
 $ lex.Xst : num 0 1 1 0 0 1 0 0 0 0 ...
 $ lex.id : int 1 2 3 4 5 6 7 8 9 10 ...
         : 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 ...
 $ dob
$ age1st : num 17.5 23.2 25.2 24.7 30.0 ...
$ agein : num 45.2 48.3 53.0 47.9 54.7 ...
 $ ageout : num 93.0 63.3 54.2 69.7 76.8 ...
 - attr(*, "time.scales")= chr "per" "age" "tfh"
 - attr(*, "breaks")=List of 3
  ..$ per: NULL
  ..$ age: NULL
  ..$ tfh: NULL
> head( nicL )
              age
                       tfh lex.dur lex.Cst lex.Xst lex.id id icd exposure
1 1934.246 45.2273 27.7465 47.7535
                                        0
                                              0
                                                        1 3
2 1934.246 48.2684 25.0820 15.0028
                                         0
                                                 1
                                                        2 4 162
                                                                        5
3 1934.246 52.9917 27.7465 1.1727
                                                        3 6 163
                                         0
                                                 1
                                                                       10
4 1934.246 47.9067 23.1861 21.7727
                                         0
                                                        4 8 527
                                                 0
                                                                        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
                                                       6 10 163
                                                                        2
                                                 1
      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).

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.Csat 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 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.

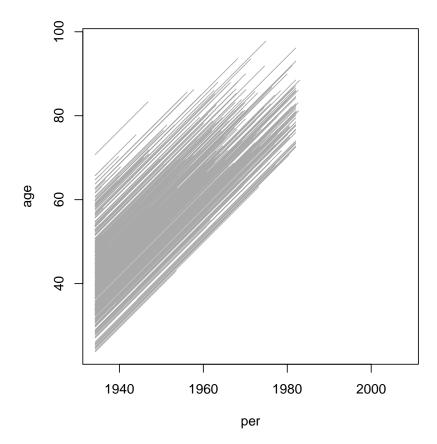


Figure 1.1: Lexis diagram of the nickel dataset.

1.1 Timescales

```
+ ylim= 10+c(0,90), yaxs="i", las=1)
> points( nicL, 1:2, pch=c(NA,3)[nicL$lex.Xst+1],
+ col="lightgray", lwd=3, cex=1.5)
> points( nicL, 1:2, pch=c(NA,3)[nicL$lex.Xst+1],
+ col=c("blue", "red")[(nicL$exp>0)+1], lwd=1, cex=1.5)
```

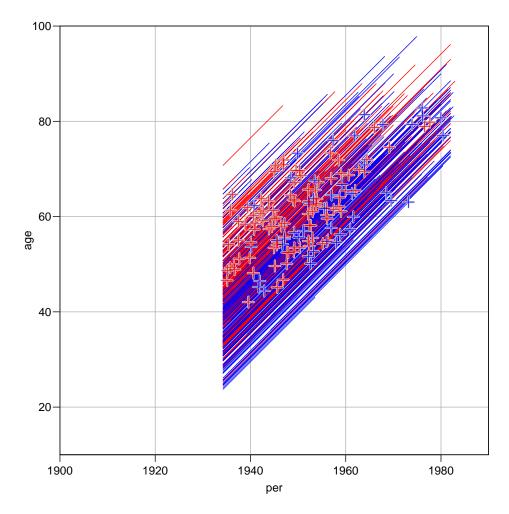


Figure 1.2: Lexis diagram of the nickel dataset, with bells and whistles. The red lines are for persons with exposure> 0, so it is pretty evident that the oldest ones are the exposed part of the cohort.

1.2 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 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>
> str( nicL )
Classes 'Lexis' and 'data.frame':
                                         679 obs. of 14 variables:
          : num 1934 1934 1934 1934 ...
 $ per
                  45.2 48.3 53.0 47.9 54.7 ...
 $ age
           : num
                  27.7 25.1 27.7 23.2 24.8 ...
 $ tfh
           : num
 $ lex.dur : num
                 47.75 15.00 1.17 21.77 22.10 ...
                  0 0 0 0 0 0 0 0 0 0 ...
 $ lex.Cst : num
                  0 1 1 0 0 1 0 0 0 0 ...
 $ lex.Xst : num
                 1 2 3 4 5 6 7 8 9 10 ...
 $ lex.id : int
          : num 3 4 6 8 9 10 15 16 17 18 ...
          : num 0 162 163 527 150 163 334 160 420 12 ...
 $ icd
 $ exposure: num 5 5 10 9 0 2 0 0.5 0 0 ...
          : num 1889 1886 1881 1886 1880 ...
 $ age1st : num 17.5 23.2 25.2 24.7 30.0 ...
          : num 45.2 48.3 53.0 47.9 54.7 ...
          : num 93.0 63.3 54.2 69.7 76.8
 - attr(*, "time.scales")= chr "per" "age" "tfh"
 - attr(*, "breaks")=List of 3
  ..$ per: NULL
  ..$ age: NULL
  ..$ tfh: NULL
> str( nicS1 )
Classes 'Lexis' and 'data.frame':
                                         2210 obs. of 14 variables:
 $ lex.id : int 1 1 1 1 1 1 2 2 2 3 ...
 $ per
          : num
                 1934 1939 1949 1959 1969 ...
 $ age
                  45.2 50.0 60.0 70.0 80.0 ...
          : num
 $ tfh
                  27.7 32.5 42.5 52.5 62.5 ...
          : num
 $ lex.dur : num
                  4.77 10.00 10.00 10.00 10.00 ...
 $ lex.Cst : num 0 0 0 0 0 0 0 0 0 0 ...
 $ lex.Xst : num
                  0 0 0 0 0 0 0 0 1 1 ...
 $ id
          : num
                  3 3 3 3 3 3 4 4 4 6 ...
          : num 0 0 0 0 0 0 162 162 163 ...
 $ exposure: num 5 5 5 5 5 5 5 5 5 10 ...
          : num 1889 1889 1889 1889 ...
 $ age1st : num 17.5 17.5 17.5 17.5 17.5 ...
 $ agein
          : num 45.2 45.2 45.2 45.2 ...
 $ ageout : num 93 93 93 93 93 ...
- attr(*, "breaks")=List of 3
  ..$ per: NULL
  ..$ age: num 0 10 20 30 40 50 60 70 80 90 ...
  ..$ tfh: NULL
 - attr(*, "time.scales")= chr "per" "age" "tfh"
> round( subset( nicS1, id %in% 8:10 ), 2 )
   lex.id
             per
                    age
                          tfh lex.dur lex.Cst lex.Xst id icd exposure
                                                                    9 1886.34
        4 1934.25 47.91 23.19
11
                                 2.09
                                                    0 8 527
                                            0
                                            0
                                                                    9 1886.34
12
        4 1936.34 50.00 25.28
                                10.00
                                                    0 8 527
13
        4 1946.34 60.00 35.28
                                 9.68
                                            0
                                                    0
                                                       8 527
                                                                    9 1886.34
        5 1934.25 54.75 24.79
                                 5.25
                                            0
                                                       9 150
                                                                    0 1879.50
14
```

24

21.29 44.33

62.54

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

The resulting object is again a Lexis object, and so follow-up may be split further along another timescale. Try this and list the result for individuals 4 and 6:

```
> nicS2 <- splitLexis( nicS1, "tfh", breaks=c(0,1,5,10,20,30,100) )
> round( subset( nicS2, id %in% 8:10 ), 2 )
```

```
tfh lex.dur lex.Cst lex.Xst id icd exposure
   lex.id
              per
                    age
                                                                            dob
13
        4 1934.25 47.91 23.19
                                  2.09
                                             0
                                                     0 8 527
                                                                      9 1886.34
                                             0
        4 1936.34 50.00 25.28
                                  4.72
                                                     0 8 527
                                                                      9 1886.34
14
        4 1941.06 54.72 30.00
                                  5.28
                                             Ω
                                                     0 8 527
                                                                      9 1886.34
15
        4 1946.34 60.00 35.28
                                  9.68
                                             0
                                                     0
                                                        8 527
                                                                      9 1886.34
16
17
        5 1934.25 54.75 24.79
                                  5.21
                                             0
                                                     Ω
                                                        9 150
                                                                      0 1879.50
18
        5 1939.46 59.96 30.00
                                  0.04
                                             0
                                                     0
                                                        9 150
                                                                      0 1879.50
19
        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 1879.50
20
                                  6.84
                                                     0 9 150
        6 1934.25 44.33 23.04
                                  5.67
                                             0
                                                                      2 1889.91
21
                                                     0 10 163
22
        6 1939.91 50.00 28.71
                                  1.29
                                             0
                                                                      2 1889.91
                                                     0 10 163
23
        6 1941.20 51.29 30.00
                                  8.71
                                             0
                                                     0 10 163
                                                                      2 1889.91
        6 1949.91 60.00 38.71
                                  2.54
                                             0
                                                     1 10 163
                                                                      2 1889.91
24
   age1st agein ageout
13
   24.72 47.91
                 69.68
   24.72 47.91
                 69.68
14
15
    24.72 47.91
                 69.68
16
    24.72 47.91
                 69.68
    29.96 54.75
17
                 76.84
18
    29.96 54.75
                 76.84
    29.96 54.75
19
                 76.84
   29.96 54.75
20
                 76.84
   21.29 44.33
                 62.54
21
   21.29 44.33 62.54
   21.29 44.33
23
                 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
                                                      45
1
                                         2.2535
                                                               25
2
           1 1936.500 47.4808 30.0000
                                         2.5192
                                                      45
                                                               65
3
                                                               65
           1 1939.019 50.0000 32.5192 10.0000
                                                      55
4
    3
           1 1949.019 60.0000 42.5192 10.0000
                                                      65
                                                               65
5
    3
           1 1959.019 70.0000 52.5192 10.0000
                                                      75
                                                               65
6
    3
           1 1969.019 80.0000 62.5192 10.0000
                                                               65
                                                      85
7
           1 1979.019 90.0000 72.5192
    3
                                         2.9808
                                                      95
                                                               65
8
    4
           2 1934.246 48.2684 25.0820
                                                      45
                                                               25
                                         1.7316
9
    4
           2 1935.978 50.0000 26.8136
                                         3.1864
                                                      55
                                                               25
10
   4
           2 1939.164 53.1864 30.0000
                                                      55
                                                               65
11
    4
           2 1945.978 60.0000 36.8136
                                         3.2712
                                                      65
                                                               65
   6
           3 1934.246 52.9917 27.7465
                                                               25
12
                                         1.1727
                                                      55
           4 1934.246 47.9067 23.1861
                                                               25
13
   8
                                         2.0933
                                                      45
14
           4 1936.340 50.0000 25.2794
                                                               25
   8
                                         4.7206
                                                      55
15
   8
           4 1941.060 54.7206 30.0000
                                         5.2794
                                                      55
                                                               65
           4 1946.340 60.0000 35.2794
                                                               65
16
                                                      65
17
   9
           5 1934.246 54.7465 24.7890
                                         5.2110
                                                      55
                                                               25
18
   9
           5 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
                                                      75
                                                               65
```

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.

1.3 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
                                                  0
                                                          4 8 527
                                                                          9
                                          0
                                                  0
                                                         5
                                                                          0
5 1934.246 54.7465 24.7890 22.0977
                                                            9 150
6 1934.246 44.3314 23.0437 18.2099
                                          0
                                                  1
                                                         6 10 163
                                                                          2
       dob 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
> nicL$agehi <- nicL$age1st + 50 / nicL$exposure</pre>
> nicC <- cutLexis( data=nicL, cut=nicL$agehi, timescale="age", new.state=2, cens=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
                                             2
                                                     2
                                                             4
                                                                8 527
                                                                             9
                                             0
                                                     0
5
     1934.246 54.7465 24.7890 22.0977
                                                             5
                                                               9 150
                                                                             0
                                             0
                                                     2
                                                             6 10 163
                                                                             2
6
     1934.246 44.3314 23.0437
                               1.9563
680
     1936.203 46.2877 25.0000 16.2536
                                             2
                                                     1
                                                             6 10 163
                                                                             2
               age1st
                        agein
                               ageout
                                          agehi
```

```
4100 1886.340 24.7206 47.9067 69.6794 30.27616
5 1879.500 29.9575 54.7465 76.8442 Inf
6 1889.915 21.2877 44.3314 62.5413 46.28770
680 1889.915 21.2877 44.3314 62.5413 46.28770
```

(The cens= 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, try:

> subset(nicS2, id %in% 8:10)

```
tfh lex.dur lex.Cst lex.Xst id icd exposure
   lex.id
               per
                       age
        4 1934.246 47.9067 23.1861
13
                                    2.0933
                                                 0
                                                          0
                                                            8 527
        4 1936.340 50.0000 25.2794
                                                            8 527
                                                                          9
14
                                    4.7206
                                                  0
                                                          0
15
        4 1941.060 54.7206 30.0000
                                    5.2794
                                                 0
                                                          Ω
                                                            8 527
                                                                          9
16
        4 1946.340 60.0000 35.2794
                                    9.6794
                                                 0
                                                         0 8 527
                                                                          9
        5 1934.246 54.7465 24.7890
                                                         0 9 150
                                                                          0
17
                                    5.2110
18
        5 1939.457 59.9575 30.0000
                                    0.0425
                                                  0
                                                         0 9 150
                                                                          0
        5 1939.500 60.0000 30.0425 10.0000
                                                         0 9 150
                                                                          0
19
                                                 0
                                                         0 9 150
        5 1949.500 70.0000 40.0425
                                                 0
                                                                          0
                                    6.8442
20
        6 1934.246 44.3314 23.0437
                                    5.6686
                                                 0
                                                         0 10 163
                                                                          2
21
22
        6 1939.915 50.0000 28.7123
                                    1.2877
                                                 0
                                                         0 10 163
                                                                          2
23
        6 1941.203 51.2877 30.0000
                                    8.7123
                                                  0
                                                          0 10 163
                                                                          2
                                                                          2
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
> nicS2$agehi <- nicS2$age1st + 50 / nicS2$exposure</pre>
> nicS2C <- cutLexis( data=nicS2, cut=nicS2$agehi, timescale="age", new.state=2, cens=0)
> subset( nicS2C, id %in% 8:10 )
                                 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
3143
          4 1936.340 50.0000 25.2794
                                     4.7206
                                                    2
                                                            2 8 527
                                                                            9
3144
          4 1941.060 54.7206 30.0000
                                      5.2794
                                                    2
                                                            2 8 527
                                                                            9
          4 1946.340 60.0000 35.2794
3145
                                      9.6794
                                                    2
                                                            2
                                                               8 527
                                                                            9
          5 1934.246 54.7465 24.7890
                                                    0
                                                              9 150
                                      5.2110
                                                                            0
18
         5 1939.457 59.9575 30.0000
                                      0.0425
                                                    0
                                                            0
                                                              9 150
                                                                            0
         5 1939.500 60.0000 30.0425 10.0000
                                                            0 9 150
19
                                                   0
                                                                            0
         5 1949.500 70.0000 40.0425
                                                   0
                                                            0 9 150
                                                                            0
20
                                      6.8442
21
         6 1934.246 44.3314 23.0437
                                      1.9563
                                                    0
                                                           2 10 163
                                                                            2
3150
         6 1936.203 46.2877 25.0000
                                      3.7123
                                                    2
                                                           2 10 163
                                                                            2
3151
          6 1939.915 50.0000 28.7123
                                      1.2877
                                                            2 10 163
3152
          6 1941.203 51.2877 30.0000
                                      8.7123
                                                            2 10 163
                                                                            2
3153
          6 1949.915 60.0000 38.7123
                                      2.5413
                                                            1 10 163
                                                                            2
          dob age1st
                        agein ageout
                                          agehi
3142 1886.340 24.7206 47.9067 69.6794 30.27616
3143 1886.340 24.7206 47.9067 69.6794 30.27616
3144 1886.340 24.7206 47.9067 69.6794 30.27616
```

```
3145 1886.340 24.7206 47.9067 69.6794 30.27616
17 1879.500 29.9575 54.7465 76.8442 Inf
18 1879.500 29.9575 54.7465 76.8442 Inf
19 1879.500 29.9575 54.7465 76.8442 Inf
20 1879.500 29.9575 54.7465 76.8442 Inf
21 1889.915 21.2877 44.3314 62.5413 46.28770
3150 1889.915 21.2877 44.3314 62.5413 46.28770
3151 1889.915 21.2877 44.3314 62.5413 46.28770
3152 1889.915 21.2877 44.3314 62.5413 46.28770
3153 1889.915 21.2877 44.3314 62.5413 46.28770
```

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 cens= argument. It names the states (in this case 0, "Alive") that will be over-ridden by new.state (in this case 2, "High exosure"). Clearly, state 1 ("Dead") should not be updated even if it is after the time where the persons moves 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.

1.4 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 a for example 1 and 2.

```
> data( nickel )
> nicL <- Lexis( entry = list( per=agein+dob,
                               age=agein,
                               tfh=agein-age1st ),
                  exit = list( age=ageout ),
           exit.status = (icd > 0)^{-} + (icd %in% c(162,163)),
                  data = nickel )
> str( nicL )
Classes 'Lexis' and 'data.frame':
                                         679 obs. of 14 variables:
          : num 1934 1934 1934 1934 ...
           : num 45.2 48.3 53.0 47.9 54.7 ...
           : num 27.7 25.1 27.7 23.2 24.8 ...
$ tfh
 $ lex.dur : num 47.75 15.00 1.17 21.77 22.10 ...
  lex.Cst : num
                  0000000000...
                  0 2 2 1 1 2 1 1 1 1 ...
  lex.Xst : int
  lex.id : int
                  1 2 3 4 5 6 7 8 9 10 ...
 $ id
                  3 4 6 8 9 10 15 16 17 18 ...
           : num
           : num 0 162 163 527 150 163 334 160 420 12 ...
 $ icd
 $ exposure: num 5 5 10 9 0 2 0 0.5 0 0 ...
          : num 1889 1886 1881 1886 1880 ...
 $ age1st : num 17.5 23.2 25.2 24.7 30.0 ...
           : num 45.2 48.3 53.0 47.9 54.7 ...
 $ ageout : num 93.0 63.3 54.2 69.7 76.8 ...
 - attr(*, "time.scales")= chr "per" "age" "tfh"
- attr(*, "breaks")=List of 3
  ..$ per: NULL
  ..$ age: NULL
  ..$ tfh: NULL
> head( nicL )
                       tfh lex.dur lex.Cst lex.Xst lex.id id icd exposure
               age
1 1934.246 45.2273 27.7465 47.7535
                                         0
                                                 0
```

```
2 1934.246 48.2684 25.0820 15.0028
                                                2
                                                       2 4 162
                                                                       5
3 1934.246 52.9917 27.7465 1.1727
                                        0
                                                2
                                                       3 6 163
                                                                      10
4 1934.246 47.9067 23.1861 21.7727
                                        0
                                                1
                                                       4 8 527
                                                                       9
5 1934.246 54.7465 24.7890 22.0977
                                                       5 9 150
                                                1
                                                                       0
6 1934.246 44.3314 23.0437 18.2099
                                                2
                                                       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
> subset( nicL, id %in% 8:10 )
      per
                      tfh lex.dur lex.Cst lex.Xst lex.id id icd exposure
              age
4 1934.246 47.9067 23.1861 21.7727
                                    0 1
                                                       4 8 527
5 1934.246 54.7465 24.7890 22.0977
                                        0
                                                1
                                                       5 9 150
                                                                       0
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:

```
> 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,
                states = c("Alive", "D.oth", "D.lung") )
> str( nicL )
Classes 'Lexis' and 'data.frame':
                                          679 obs. of 14 variables:
          : num 1934 1934 1934 1934 ...
 $ per
 $ age
           : num 45.2 48.3 53.0 47.9 54.7 ...
          : num 27.7 25.1 27.7 23.2 24.8 ...
 $ tfh
 $ lex.dur : num 47.75 15.00 1.17 21.77 22.10 ...
 $ lex.Cst : Factor w/ 3 levels "Alive", "D.oth", ...: 1 1 1 1 1 1 1 1 1 1 ...
 $ lex.Xst : Factor w/ 3 levels "Alive", "D.oth",..: 1 2 2 3 3 2 3 3 3 ...
 $ lex.id : int 1 2 3 4 5 6 7 8 9 10 ...
                  3 4 6 8 9 10 15 16 17 18 ...
 $ id
           : num
 $ 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 ...
 $ age1st : num 17.5 23.2 25.2 24.7 30.0 ...
          : num 45.2 48.3 53.0 47.9 54.7 ...
 $ ageout : num 93.0 63.3 54.2 69.7 76.8 ...
 - attr(*, "time.scales")= chr "per" "age" "tfh"
- attr(*, "breaks")=List of 3
  ..$ per: NULL
  ..$ age: NULL
  ..$ tfh: NULL
```

You can get an overview of the number of records by state and transitions between states as well as the perosn-years in each state by using tab.Lexis(), try:

> tab.Lexis(nicL)

```
States:
     #records:
        Alive D.oth D.lung Sum
From
                                 #events:
                                           #risk time:
                                                              Rate
                                                                          (95%
 Alive
           47
                137
                        495 679
                                      632
                                              15348.06 0.04117785 0.03808940
States:
     #records:
From
           c.int.)
  Alive 0.04451673
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 it was
in a state we
> nicL$agehi <- nicL$age1st + 50 / nicL$exposure</pre>
> nicC <- cutLexis( data=nicL, cut=nicL$agehi, "age", new.state="HiExp", cens="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
                                                                8 527
                                         HiExp
                                                D.lung
                                                             4
     1934.246 54.7465 24.7890 22.0977
                                                             5 9 150
                                                                              0
                                         Alive
                                                 D.lung
     1934.246 44.3314 23.0437
                               1.9563
                                         Alive
                                                  HiExp
                                                             6 10 163
                                                                              2
680
    1936.203 46.2877 25.0000 16.2536
                                                  D.oth
                                                             6 10 163
                                                                              2
                                         HiExp
              age1st
                        agein ageout
4100 1886.340 24.7206 47.9067 69.6794 30.27616
     1879.500 29.9575 54.7465 76.8442
6
     1889.915 21.2877 44.3314 62.5413 46.28770
    1889.915 21.2877 44.3314 62.5413 46.28770
> tab.Lexis( nicC )
States:
     #records:
From
        Alive D.oth D.lung HiExp Sum
                                       #events:
                                                  #risk time:
                                                                     Rate
                 65
                        279
                                                    10772.533 0.03963785
 Alive
           39
                               83 466
                                            427
 HiExp
                 72
                               8 296
                                            288
            0
                        216
                                                     4575.524 0.06294361
 Sum
           39
                137
                        495
                               91 762
                                            715
                                                    15348.057 0.04658570
States:
     #records:
     To
              (95%
From
                       c.int.)
  Alive 0.03605095 0.04358162
 HiExp 0.05607809 0.07064967
        0.04329312 0.05012869
```

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.

1.5 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 cens are ignored.