# Follow-up data with the Epi package

Summer 2014

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	7
5	Competing risks — multiple types of events	10
6	Multiple events of the same type (recurrent events)	13

```
> library(Epi)
> print( sessionInfo(), 1=F )
```

R version 3.4.0 (2017-04-21)

Platform: x86\_64-pc-linux-gnu (64-bit) Running under: Ubuntu 14.04.5 LTS

Matrix products: default

BLAS: /usr/lib/libblas/libblas.so.3.0 LAPACK: /usr/lib/lapack/liblapack.so.3.0

attached base packages:

[1] stats graphics grDevices utils datasets methods base

other attached packages:

[1] Epi\_2.15

loaded via a namespace (and not attached):

[1]	cmprsk_2.2-7	MASS_7.3-45	compiler_3.4.0	Matrix_1.2-6
[5]	plyr_1.8.4	parallel_3.4.0	tools_3.4.0	survival_2.41-3
[9]	etm_0.6-2	Rcpp_0.12.5	splines_3.4.0	grid_3.4.0
F407	D : 0044 0 4			

[13] numDeriv\_2014.2-1 lattice\_0.20-33

## 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").

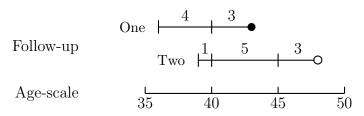


Figure 1: Follow-up of two persons

### 2 Timescales

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

2 Timescales

- 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 (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
                  3 4 6 8 9 10 15 16 17 18 ...
           : num
$ icd
                  0 162 163 527 150 163 334 160 420 12 ...
           : num
$ exposure: num 5 5 10 9 0 2 0 0.5 0 0 ...
$ dob
           : num
                 1889 1886 1881 1886 1880 ...
                  17.5 23.2 25.2 24.7 30 ...
$ age1st
           : num
$ agein
                 45.2 48.3 53 47.9 54.7 ...
           : num
$ ageout
                 93 63.3 54.2 69.7 76.8 ...
           : num
```

#### > str( nicL )

```
Classes âĂŸLexisâĂŹ and 'data.frame':
                                              679 obs. of 14 variables:
                  1934 1934 1934 1934 . . .
           : num
 $ age
           : num
                  45.2 48.3 53 47.9 54.7 ...
                  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 ...
 $ 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 ...
 $ id
           : 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 ...
                 1889 1886 1881 1886 1880 ...
 $ dob
           : num
 $ age1st : num 17.5 23.2 25.2 24.7 30 ...
                 45.2 48.3 53 47.9 54.7 ...
 $ agein
           : num
 $ ageout : num 93 63.3 54.2 69.7 76.8 ...
 - attr(*, "time.scales")= chr
                                "per" "age" "tfh"
                                0.01 \quad 0.01 \quad 0.01
 - attr(*, "time.since")= chr
 - 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
                                                          1
                                                             3
                                                                 0
                                                                          5
2 1934.246 48.2684 25.0820 15.0028
                                          0
                                                  1
                                                          2
                                                             4 162
                                                                          5
                                                          3
3 1934.246 52.9917 27.7465
                                          0
                                                  1
                                                             6 163
                                                                         10
                           1.1727
4 1934.246 47.9067 23.1861 21.7727
                                          0
                                                  0
                                                          4
                                                             8 527
                                                                          9
                                                  0
5 1934.246 54.7465 24.7890 22.0977
                                          0
                                                          5
                                                             9 150
                                                                          0
6 1934.246 44.3314 23.0437 18.2099
                                                  1
                                                                          2
                                          0
                                                          6 10 163
            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.

# 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) )
> summary( nicL )

Transitions:
    To
From 0 1 Records: Events: Risk time: Persons:
    0 542 137 679 137 15348.06 679
```

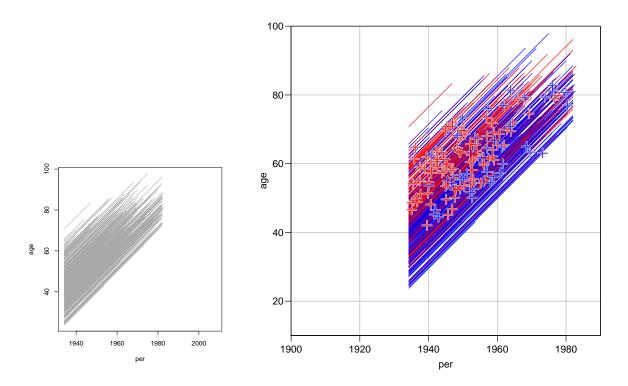


Figure 2: Lexis diagram of the **nickel** dataset, left panel the default version, the right one with bells and whistles. The red lines are for persons with exposure> 0, so it is pretty evident that the oldest ones are the exposed part of the cohort.

#### > summary( nicS1 )

#### Transitions:

То

From 0 1 Records: Events: Risk time: Persons: 0 2073 137 2210 137 15348.06 679

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:

#### > round( subset( nicS1, id %in% 8:10 ), 2 )

	lex.id	per	age	tfh	lex.dur	lex.Cst	lex.Xst	id	icd	exposure	dob
11	4	1934.25	47.91	23.19	2.09	0	0	8	527	9	1886.34
12	4	1936.34	50.00	25.28	10.00	0	0	8	527	9	1886.34
13	4	1946.34	60.00	35.28	9.68	0	0	8	527	9	1886.34
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
16	5	1949.50	70.00	40.04	6.84	0	0	9	150	0	1879.50
17	6	1934.25	44.33	23.04	5.67	0	0	10	163	2	1889.91

```
0
18
        6 1939.91 50.00 28.71
                                  10.00
                                                        0 10 163
                                                                         2 1889.91
        6 1949.91 60.00 38.71
                                   2.54
                                               0
19
                                                        1 10 163
                                                                         2 1889.91
   age1st agein ageout
11
    24.72 47.91
                  69.68
    24.72 47.91
                  69.68
12
13
    24.72 47.91
                  69.68
    29.96 54.75
14
                  76.84
    29.96 54.75
                  76.84
15
    29.96 54.75
                  76.84
16
    21.29 44.33
                  62.54
17
18
    21.29 44.33
                  62.54
    21.29 44.33
                  62.54
19
```

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) )
> round( subset( nicS2, id %in% 8:10 ), 2 )
```

	<pre>lex.id</pre>	per	age	tfh	${\tt lex.dur}$	${\tt lex.Cst}$	lex.Xst	id	icd	exposure	dob
13	4	1934.25	47.91	23.19	2.09	0	0	8	527	9	1886.34
14	4	1936.34	50.00	25.28	4.72	0	0	8	527	9	1886.34
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
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
20	5	1949.50	70.00	40.04	6.84	0	0	9	150	0	1879.50
21	6	1934.25	44.33	23.04	5.67	0	0	10	163	2	1889.91
22	6	1939.91	50.00	28.71	1.29	0	0	10	163	2	1889.91
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	0	1	10	163	2	1889.91

age1st agein ageout

```
13 24.72 47.91 69.68
```

<sup>14 24.72 47.91 69.68</sup> 

<sup>15 24.72 47.91 69.68</sup> 

<sup>16 24.72 47.91 69.68</sup> 

<sup>17 29.96 54.75 76.84</sup> 

<sup>18 29.96 54.75 76.84</sup> 19 29.96 54.75 76.84

<sup>-- -- -- -- -- -- --</sup>

<sup>20 29.96 54.75 76.84</sup> 21 21.29 44.33 62.54

<sup>22 21.29 44.33 62.54</sup> 

<sup>23 21.29 44.33 62.54</sup> 

<sup>24 21.29 44.33 62.54</sup> 

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:

```
> timeBand( nicS2, "age", "middle" )[1:20]
 [1] 45 45 55 65 75 85 95 45 55 55 65 55 45 55 65 55 65 75
> # For nice printing and column labelling use the data.frame() function:
 data.frame( nicS2[,c("id","lex.id","per","age","tfh","lex.dur")],
               mid.age=timeBand( nicS2, "age", "middle" ),
               mid.tfh=timeBand( nicS2, "tfh", "middle" ) )[1:20,]
   id lex.id
                                    tfh lex.dur mid.age mid.tfh
                   per
                           age
    3
           1 1934.246 45.2273 27.7465
                                         2.2535
1
                                                      45
                                                              25
2
    3
           1 1936.500 47.4808 30.0000
                                                      45
                                                              65
                                         2.5192
3
    3
           1 1939.019 50.0000 32.5192 10.0000
                                                      55
                                                              65
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
                                                      85
                                                              65
7
    3
           1 1979.019 90.0000 72.5192
                                         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
                                                      55
                                                              25
                                         3.1864
10
    4
           2 1939.164 53.1864 30.0000
                                                      55
                                         6.8136
                                                              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
    8
           4 1936.340 50.0000 25.2794
                                         4.7206
                                                      55
                                                              25
14
    8
           4 1941.060 54.7206 30.0000
15
                                         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
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
    9
           5 1949.500 70.0000 40.0425
                                         6.8442
                                                      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.

# 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
       per
4 1934.246 47.9067 23.1861 21.7727
                                                             8 527
                                          0
                                                  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
                                                   1
                                                          6 10 163
                                                                          2
                                          0
       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
> agehi <- nicL$age1st + 50 / nicL$exposure</pre>
> nicC <- cutLexis( data=nicL, cut=agehi, timescale="age",
                     new.state=2, precursor.states=0 )
> subset( nicC, id %in% 8:10 )
                          tfh lex.dur lex.Cst lex.Xst lex.id id icd exposure
683 1934.246 47.9067 23.1861 21.7727
                                            2
                                                     2
                                                               8 527
                                                                            0
    1934.246 54.7465 24.7890 22.0977
                                            0
                                                    0
                                                               9 150
                                                            5
    1934.246 44.3314 23.0437
                                            0
                                                     2
                                                            6 10 163
                                                                            2
                               1.9563
                                            2
685 1936.203 46.2877 25.0000 16.2536
                                                     1
                                                            6 10 163
                                                                            2
         dob
             age1st
                        agein
                               ageout
683 1886.340 24.7206 47.9067 69.6794
    1879.500 29.9575 54.7465 76.8442
    1889.915 21.2877 44.3314 62.5413
685 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	per	age	tfh	lex.dur	${\tt lex.Cst}$	lex.Xst	${\tt id}$	icd	exposure
13	4	1934.246	47.9067	23.1861	2.0933	0	0	8	527	9
14	4	1936.340	50.0000	25.2794	4.7206	0	0	8	527	9
15	4	1941.060	54.7206	30.0000	5.2794	0	0	8	527	9
16	4	1946.340	60.0000	35.2794	9.6794	0	0	8	527	9
17	5	1934.246	54.7465	24.7890	5.2110	0	0	9	150	0
18	5	1939.457	59.9575	30.0000	0.0425	0	0	9	150	0
19	5	1939.500	60.0000	30.0425	10.0000	0	0	9	150	0
20	5	1949.500	70.0000	40.0425	6.8442	0	0	9	150	0
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	2

```
23
        6 1941.203 51.2877 30.0000 8.7123
                                                  0
                                                          0 10 163
                                                                          2
        6 1949.915 60.0000 38.7123 2.5413
                                                          1 10 163
24
                                                  0
                                                                          2
        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
> 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
                                                            2 8 527
                                                                            9
          4 1936.340 50.0000 25.2794
                                                    2
                                                            2
                                                               8 527
                                                                            9
3143
                                      4.7206
          4 1941.060 54.7206 30.0000
                                                    2
                                                            2 8 527
                                                                             9
3144
                                     5.2794
          4 1946.340 60.0000 35.2794
                                                    2
                                                            2 8 527
                                                                             9
3145
                                      9.6794
17
          5 1934.246 54.7465 24.7890
                                      5.2110
                                                    0
                                                               9 150
                                                                             0
18
          5 1939.457 59.9575 30.0000
                                      0.0425
                                                    0
                                                            0
                                                               9 150
                                                                             0
19
          5 1939.500 60.0000 30.0425 10.0000
                                                    0
                                                            0
                                                              9 150
                                                                             0
20
          5 1949.500 70.0000 40.0425 6.8442
                                                    0
                                                            0 9 150
                                                                             0
          6 1934.246 44.3314 23.0437
                                                                             2
21
                                                    0
                                                            2 10 163
                                      1.9563
          6 1936.203 46.2877 25.0000 3.7123
                                                    2
                                                                            2
3150
                                                            2 10 163
          6 1939.915 50.0000 28.7123
                                                    2
                                                            2 10 163
                                                                             2
3151
                                      1.2877
3152
          6 1941.203 51.2877 30.0000
                                      8.7123
                                                    2
                                                            2 10 163
                                                                             2
          6 1949.915 60.0000 38.7123
                                                    2
                                                                             2
3153
                                      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
     1879.500 29.9575 54.7465 76.8442
18
19
     1879.500 29.9575 54.7465 76.8442
     1879.500 29.9575 54.7465 76.8442
20
     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
                 Records:
                            Events: Risk time:
   0 47 495 137
                       679
                                632
                                      15348.06
                                                      679
> subset( nicL, id %in% 8:10 )
                       tfh lex.dur lex.Cst lex.Xst lex.id id icd exposure
       per
               age
4 1934.246 47.9067 23.1861 21.7727
                                          0
                                                  1
                                                             8 527
5 1934.246 54.7465 24.7890 22.0977
                                          0
                                                   1
                                                          5 9 150
                                                                          0
                                                  2
6 1934.246 44.3314 23.0437 18.2099
                                          0
                                                          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:

39

0

39

Alive

HiExp

Sum

279

216

495

65

72

137

83

91

8

```
> 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") )
> summary( nicL )
Transitions:
     To
From
        Alive D.oth D.lung Records:
                                       Events: Risk time:
           47
                495
                        137
                                           632
                                                  15348.06
  Alive
                                  679
                                                                 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 )
                 age
                         tfh lex.dur lex.Cst lex.Xst lex.id id icd exposure
         per
                                                              8 527
683 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
                                                           6 10 163
                                                                            2
                              1.9563
                                        Alive
                                                HiExp
685 1936.203 46.2877 25.0000 16.2536
                                        HiExp D.lung
                                                           6 10 163
                                                                            2
         dob
              age1st
                       agein
                               ageout
                                         agehi
683 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
685 1889.915 21.2877 44.3314 62.5413 46.28770
> summary( nicC, scale=1000 )
Transitions:
     Tο
                                             Events: Risk time:
        Alive HiExp D.oth D.lung
                                  Records:
From
```

427

288

715

10.77

15.35

4.58

466

296

679

466

296

762

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:

```
> nicC <- cutLexis( data = nicL,
                      cut = nicL$agehi,
                timescale = "age",
+
                new.state = "Hi",
+
              split.states=TRUE, new.scale=TRUE,
        precursor.states = "Alive" )
 subset( nicC, id %in% 8:10 )
                                                            lex.Xst lex.id id icd
                          tfh
                                Hi.dur lex.dur lex.Cst
         per
                  age
683 1934.246 47.9067 23.1861 17.63054 21.7727
                                                     Ηi
                                                         D.oth(Hi)
                                                                         4
                                                                           8 527
    1934.246 54.7465 24.7890
                                    NA 22.0977
                                                  Alive
                                                              D.oth
                                                                            9 150
    1934.246 44.3314 23.0437
                                         1.9563
                                                  Alive
                                                                 Ηi
                                                                         6 10 163
                               0.00000 16.2536
                                                     Hi D.lung(Hi)
                                                                         6 10 163
685 1936.203 46.2877 25.0000
                   dob
    exposure
                        age1st
                                 agein
                                         ageout
                                                   agehi
683
           9 1886.340 24.7206 47.9067 69.6794 30.27616
5
           0 1879.500 29.9575 54.7465 76.8442
6
           2 1889.915 21.2877 44.3314 62.5413 46.28770
685
           2 1889.915 21.2877 44.3314 62.5413 46.28770
> summary( nicC, scale=1000 )
Transitions:
     To
        Alive Hi D.oth D.lung D.lung(Hi) D.oth(Hi)
                                                      Records:
From
           39 83
                                         0
  Alive
                    279
                            65
                                                   0
                                                            466
                                                                     427
  Ηi
            0
                      0
                             0
                                        72
                                                 216
                                                            296
                                                                     288
```

72

216

762

715

#### Transitions:

Sum

То

From Risk time: Persons:

39 91

279

65

Alive	10.77	466
Hi	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.