Model B: Estimation Results

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Estimation results of **Model B**.

Load environmet

```
base::source("http://www.ucl.ac.uk/~ucakadl/ELECT/ELECT.r") # load ELECT functions
base::source("./scripts/ELECT-utility-functions.R") # ELECT utility functions
library(magrittr) #Pipes
library(msm)
requireNamespace("ggplot2", quietly=TRUE)
requireNamespace("dplyr", quietly=TRUE)
requireNamespace("testit", quietly=TRUE)
```

Save fitted models here :

[1] "./data/shared/derived/models/model-b/"

Load data

```
# first, the script `O-ellis-island.R` imports and cleans the raw data
# second, the script `1-encode-multistate.R` augments the data with multi-states
# load this data transfer object (dto)
dto <- readRDS("./data/unshared/derived/dto.rds")</pre>
names(dto)
names(dto[["unitData"]])
                              # 1st element - unit(person) level data
names(dto[["metaData"]])
                              # 2nd element - meta data, info about variables
names(dto[["ms_mmse"]])
                             # 3rd element - data for MMSE outcome
ds_miss <- dto$ms_mmse$missing # data after encoding missing states (-1, -2)
ds_ms <- dto$ms_mmse$multi</pre>
                             # data after encoding multistates (1,2,3,4)
# compare before and after ms encoding
view_id <- function(ds1,ds2,id){</pre>
  cat("Before ms encoding:","\n")
 print(ds1[ds1$id==id,])
  cat("\nAfter ms encoding","\n")
  print(ds2[ds2$id==id,])
ids <- sample(unique(ds_miss$id),1) # view a random person for sporadic inspections
# ids <- c(75507759) #, 37125649, 50101073, 6804844, 83001827 , 56751351, 13485298, 56751351, 75507759)
view_id(ds_miss, ds_ms, ids)
Before ms encoding:
           id fu_year died age_bl male edu age_death age_at_visit mmse presumed_alive
6318 59166782
                   0
                        0 83.8412 FALSE 16
                                                          83.84120
                                                                                   TRUE
                                                   NA
                                                                     29
                        0 83.8412 FALSE 16
                                                                                   TRUE
6319 59166782
                   1
                                                   NA
                                                          84.84052
                                                                      30
6320 59166782
                   2 0 83.8412 FALSE 16
                                                   NA
                                                           85.81793
                                                                     29
                                                                                   TRUE
6321 59166782
                   3 0 83.8412 FALSE 16
                                                   NA
                                                           86.84736
                                                                     -2
                                                                                   TRUE
After ms encoding
           id fu_year died age_bl male edu
                                                 age state presumed_alive mmse firstobs
6318 59166782
                0
                        0 83.8412 FALSE 16 83.84120
                                                         1
                                                                     TRUE
                                                                            29
                                                                                       1
6319 59166782
                   1
                        0 83.8412 FALSE 16 84.84052
                                                                     TRUE
                                                                            30
                                                                                       0
                                                         1
                   2 0 83.8412 FALSE 16 85.81793
6320 59166782
                                                                      TRUE
                                                                             29
                                                                                       0
                                                         1
                                                        -2
6321 59166782
                   3 0 83.8412 FALSE 16 86.84736
                                                                      TRUE
                                                                            -2
```

Remove cases

```
## REMOVE observations with missing age
cat("\n Number of observations with missing age :", sum(is.na(ds_ms$age)),"\n")

Number of observations with missing age : 1

ds_clean <- ds_ms %>%
   dplyr::filter(!is.na(age))
cat("\n Number of observations with missing age :", sum(is.na(ds_clean$age)),"\n")
```

Number of observations with missing age : 0

```
## REMOVE subjects with only ONE observed data point
cat("\n Count how many subjects have *n* observed data points")
Count how many subjects have *n* observed data points
# ds miss %>%
ds_clean %>%
  dplyr::group_by(id) %>%
  dplyr::summarize(n_data_points = n()) %>%
  dplyr::group_by(n_data_points) %>%
  dplyr::summarize(n_people=n()) %>%
  print()
# A tibble: 17 \times 2
   n_{data\_points} n_{people}
           <int>
                    <int>
                      119
1
               1
               2
2
                      205
3
               3
                      184
               4
4
                      180
5
               5
                      190
6
               6
                      104
7
               7
                      108
8
               8
                      113
9
               9
                      127
              10
                      116
10
                      110
              11
11
12
              12
                       71
13
              13
                       21
14
              14
                       14
15
              15
                       13
16
              16
                       17
                        3
17
              17
# Determine which ids have only a single observation
remove_ids <- ds_clean %>%
  dplyr::group_by(id) %>%
  dplyr::summarize(n_data_points = n()) %>%
  dplyr::arrange(n_data_points) %>%
  dplyr::filter(n_data_points==1) %>%
  dplyr::select(id)
remove_ids <- remove_ids$id</pre>
cat("\n The number of subjects to be removed :",length(remove_ids) )
 The number of subjects to be removed: 119
ds clean <- ds clean %>%
  dplyr::filter(!(id %in% remove_ids))
```

Count how many subjects have *n* observed data points

cat("\n Count how many subjects have *n* observed data points")

```
# ds_miss %>%
ds_clean %>%
  dplyr::group_by(id) %>%
  dplyr::summarize(n_data_points = n()) %>%
  dplyr::group_by(n_data_points) %>%
  dplyr::summarize(n_people=n()) %>%
  print()
# A tibble: 16 \times 2
   n_data_points n_people
           <int>
                    <int>
               2
                       205
2
               3
                       184
                4
3
                       180
               5
4
                       190
               6
5
                       104
6
               7
                       108
7
               8
                       113
               9
                       127
8
9
               10
                       116
10
                       110
               11
11
               12
                        71
12
               13
                        21
               14
                        14
13
14
               15
                        13
15
               16
                        17
16
               17
                         3
## REMOVE subjects with IMS at firstobs = TRUE
cat("\n View subjects with intermediate missing state at first observation")
 View subjects with intermediate missing state at first observation
ids_firstobs_ims <- ds_clean %>%
  dplyr::filter(firstobs == TRUE & state == -1) %>%
  dplyr::select(id) %>% print()
1 80333458
2 90214403
3 90447310
4 91804757
ids_firstobs_ims <- ids_firstobs_ims[,"id"]</pre>
ds_clean <- ds_clean %>%
  dplyr::filter(!id %in% ids_firstobs_ims)
cat("\n View subjects with intermediate missing state at first observation")
\label{thm:linear_variance} \mbox{\sc View subjects with intermediate missing state at first observation}
ds clean %>%
  dplyr::filter(firstobs == TRUE & state == -1) %>%
  dplyr::select(id) %>% print()
```

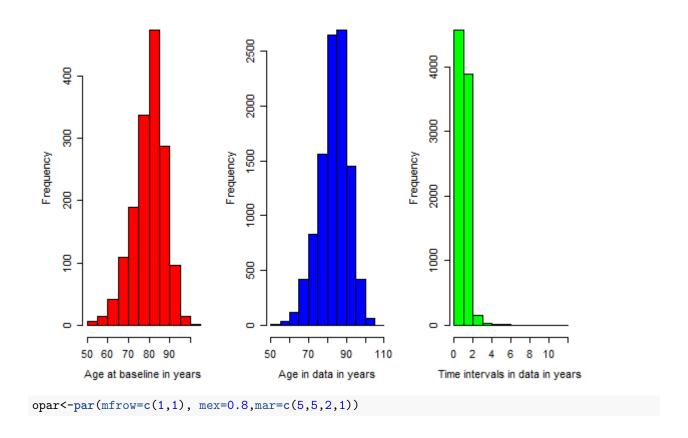
```
<0 rows> (or 0-length row.names)
```

Categorize covariates

Age diagnostic

```
# Time intervals in data:
# the age difference between timepoint for each individual
intervals <- matrix(NA,nrow(ds_clean),2)</pre>
for(i in 2:nrow(ds_clean)){
  if(ds_clean$id[i] == ds_clean$id[i-1]){
    intervals[i,1] <- ds_clean$id[i]</pre>
    intervals[i,2] <- ds_clean$age[i]-ds_clean$age[i-1]</pre>
 }
  intervals <- as.data.frame(intervals)</pre>
  colnames(intervals) <- c("id", "interval")</pre>
cat("\n Minimum interval length : ",min(intervals, na.rm=T))
Minimum interval length: 0.00273785
cat("\n Maximum interval length : ", max(intervals, na.rm=T))
Maximum interval length: 99982430
# the age difference between timepoint for each individual
intervals <- intervals[!is.na(intervals[,2]),] # Remove NAs:</pre>
cat("\nTime intervals between observations within individuals:\n")
Time intervals between observations within individuals:
print(round(quantile(intervals[,2]),digits))
        25%
              50%
                    75% 100%
0.00 0.96 1.00 1.03 11.87
# Info on age and time between observations:
opar<-par(mfrow=c(1,3), mex=0.8,mar=c(5,5,3,1))
```

```
hist(ds_clean$age[ds_clean$firstobs==1],col="red",xlab="Age at baseline in years",main="")
hist(ds_clean$age,col="blue",xlab="Age in data in years",main="")
hist(intervals[,2],col="green",xlab="Time intervals in data in years",main="")
```



Estimation prep

Number of subjects with either IMS or RC state(s): 149

```
ids_with_both <- dplyr::intersect(ids_with_im, ids_with_rc)</pre>
cat("\n Number of subjects with both IMS and RC state(s) : ",length(ids_with_both) )
Number of subjects with both IMS and RC state(s): 1
# subset a random sample of individuals if needed
set.seed(42)
ids <- sample(unique(ds clean$id), 100)
# define the data object to be passed to the estimation call
ds <- ds_clean %>%
 # dplyr::filter(id %in% ids) %>% # make sample smaller if needed
 # exclude individuals with missing states
 # dplyr::filter(!id %in% ids_with_im) %>%
 # dplyr::filter(!id %in% ids_with_rc) %>%
 dplyr::mutate(
   male = as.numeric(male),
   age = (age - 75), # centering
   age_bl = (age_bl - 75) # centering
) %>%
 dplyr::select(id, age_bl,male, edu, educat, educatF,firstobs, fu_year, age, state)
# view data object to be passed to the estimation call
set.seed(42)
ids <- sample(unique(ds$id), 1)</pre>
ds %>% dplyr::filter(id %in% ids)
                                     educatF firstobs fu_year
             age_bl male edu educat
                                                                    age state
1 90544686 7.696783
                      0 12
                                                           0 7.696783
                                 1 >11 years
                                                   1
2 90544686 7.696783
                       0 12
                                  1 >11 years
                                                    0
                                                           1 8.682409
3 90544686 7.696783
                       0 12
                                 1 >11 years
                                                   0
                                                           2 9.731006
4 90544686 7.696783
                    0 12
                                 1 >11 years
                                                   0
                                                           3 10.689254
                     0 12
                                                   0
5 90544686 7.696783
                                 1 >11 years
                                                           4 11.691307
                                                  0
                                                          5 12.709788
6 90544686 7.696783
                    0 12
                                 1 >11 years
                                                                           1
                                                  0
7 90544686 7.696783 0 12
                                 1 >11 years
                                                          6 13.665298
                                                           7 14.678303
8 90544686 7.696783
                    0 12
                                                    0
                                                                           2
                                  1 >11 years
9 90544686 7.696783
                       0 12
                                  1 >11 years
                                                    0
                                                           8 15.680356
                                                                            1
10 90544686 7.696783
                       0 12
                                  1 >11 years
                                                    0
                                                            9 16.709788
                                                                            1
cat("\n Subject count : ",length(unique(ds$id)),"\n")
Subject count: 1572
cat("\n Frequency of states at baseline\n")
Frequency of states at baseline
sf <- ds %>%
 dplyr::filter(firstobs==TRUE) %>%
 dplyr::group_by(state) %>%
 dplyr::summarize(count = n()) %>% # basic frequiencies
 dplyr::mutate(pct = round(count/sum(count),2)) %%  # percentages, use for starter values
 print()
# A tibble: 3 \times 3
```

state count pct

```
<dbl> <int> <dbl>
     1 1189 0.76
1
2
         281 0.18
3
         102 0.06
      3
cat("\nState table: \n")
State table:
print(msm::statetable.msm(state,id,data=ds)) # transition frequencies
      -2
                      2
                            3
from
            -1
                 1
  -2
      32
            0
                 0
                      0
  -1
       0
           25
                27
                     13
                          26
                               47
           59 4855 715 120 251
  1
      32
  2
       8
           20 534
                    478 256 146
           34
                24
                     96 649 232
# these will be passed as starting values
initial_probabilities <- as.numeric(as.data.frame(sf[!sf$state %in% c(-1,-2),"pct"])$pct)</pre>
initial_probabilities <- c(initial_probabilities,0) # no death state at first observation
cat('\n The inital values for estimation : ', initial_probabilities)
The inital values for estimation: 0.76 0.18 0.06 0
# save the object to be used during estimation
saveRDS(ds, "./data/unshared/ds_estimation.rds")
```

Specifications

Fitting function

```
estimate_multistate <- function(</pre>
 model name
  ,ds
                        # data object
  ,Q
                        # Q-matrix of transitions
  ,E
                        # misspecification matrix
  ,qnames
                        # names of the rows in the Q matrix
  ,cov_names
                        # string with covariate names
  covariates_ <- as.formula(paste0("~",cov_names))</pre>
 model <- msm(
   formula
                 = state ~ age,
   subject
                 = id,
   data
                  = ds,
   center
                 = FALSE,
   qmatrix
                  = Q,
                 = E,
   ematrix
   death
                 = TRUE,
    covariates = covariates_,
    censor = c(-1,-2),
```

```
censor.states = list(c(1,2,3), c(1,2,3)),
    method = method_,
    constraint = constraint_,
    fixedpars = fixedpars_,
initprobs = initprobs_,
    est.initprobs = TRUE,
               = list(trace=0,REPORT=1,maxit=1000,fnscale=10000)
    control
  # model <- pasteO("test", covariates_)</pre>
  saveRDS(model, paste0(pathSaveFolder,model_name,".rds"))
  return(model)
}
msm_summary <- function(model){</pre>
cat("\n-2loglik =", model$minus2loglik,"\n")
cat("Convergence code =", model$opt$convergence,"\n")
     <- model $ opt $ par
p.se <- sqrt(diag(solve(1/2*model$opt$hessian)))</pre>
print(cbind(p=round(p,digits),
            se=round(p.se,digits),"Wald ChiSq"=round((p/p.se)^2,digits),
            "Pr>ChiSq"=round(1-pchisq((p/p.se)^2,df=1),digits)),
      quote=FALSE)
}
```

Model

```
q < - .01
# transition matrix
Q \leftarrow rbind(c(0, q, 0, q),
           c(q, 0, q, q),
           c(0, 0, 0, q),
           c(0, 0, 0, 0)
# misclassification matrix
E \leftarrow rbind(c(0, 0, 0, 0),
           c(0, 0, .1, 0),
           c(0, 0, 0, 0),
           c(0, 0, 0, 0)
# transition names
qnames = c(
 "Healthy - Mild", # q12
 # "Healthy - Severe", # q13
 "Healthy - Dead", # q14
 "Mild - Healthy", # q21
 "Mild - Severe", # q23
 "Mild - Dead",
                  # 924
 # "Severe - Healthy",# q31
 # "Severe - Mild", # q32
 "Severe - Dead" # q34
)
```

msm options

```
digits = 2
method_ = "BFGS"
                      # alternatively, if does not converge "Nedler-Mead"
constraint_ = NULL
                      # additional model constraints
fixedpars_ = NULL
                      # fixed parameters
initprobs_ = initial_probabilities
# turn this chunk OFF when printing the report
# compile model objects with msm() call
# each model will be saved in the specified folder, namely pathSaveFolder
# estimate_multistate("mA1", ds, Q, E, qnames,cov_names = "age")
# estimate_multistate("mA2", ds, Q, E, qnames,cov_names = "age + age_bl")
# estimate_multistate("mA3", ds, Q, E, qnames,cov_names = "age + age_bl + male")
# estimate_multistate("mA4", ds, Q, E, qrames,cov_names = "age + age_bl + male + educat")
# estimate_multistate("mA5", ds, Q, E, qnames,cov_names = "age + age_bl + male + edu")
# assemble the list object with the results of msm estimation
models <- list()
models[["age"]][["msm"]] <- readRDS(paste0(pathSaveFolder, 'mA1.rds'))</pre>
models[["age_bl"]][["msm"]] <- readRDS(paste0(pathSaveFolder,'mA2.rds'))</pre>
models[["male"]][["msm"]] <- readRDS(paste0(pathSaveFolder,'mA3.rds'))</pre>
models[["educat"]][["msm"]] <- readRDS(paste0(pathSaveFolder,'mA4.rds'))</pre>
models[["edu"]][["msm"]] <- readRDS(paste0(pathSaveFolder, 'mA5.rds'))</pre>
```

elect options

```
alive_states <- c(1,2,3)
ds_alive <- ds[ds$state %in% alive_states,]</pre>
fixedpars <- fixedpars_</pre>
age_min <- 0
age_max < -35
age_bl <- 0
male <-0
educat <- 0
edu <- 11
replication n <- 1000
time_scale <- "years"</pre>
grid_par <- .5</pre>
# turn this chunk OFF when printing the report
# for(model_ in names(models) ){
   # determine covariate list
  if(model_=="age"){covar_list = list(age=age_min)}
#
   if(model_=="aqe_bl"){covar_list = list(aqe=aqe_min, aqe_bl=aqe_bl)}
   if(model_=="male"){covar_list = list(age=age_min, age_bl=age_bl, male=male)}
#
   if(model_=="educat"){covar_list = list(aqe=aqe_min, aqe_bl=aqe_bl, male=male, educat=educat)}
#
   if(model_=="edu")\{covar\_list = list(age=age\_min, age\_bl=age\_bl, male=male, edu=edu)\}
#
   # compute LE
# models[[model_]][["LE"]] <- elect(</pre>
#
    model = models[[model]][["msm"]], # fitted msm model
      b.covariates = covar_list, # list with specified covarites values
```

```
statedistdata = ds_alive, # data for distribution of living states
      time.scale.msm = time_scale, # time scale in multi-state model ("years", ...)
#
#
                    = grid_par, # grid parameter for integration
#
                     = age_max, # assumed maximum age in years
      age.max
#
      S
                     = replication_n # number of simulation cycles
    )
#
#
    # models[[model_]][["LE"]] <- models[["msm"]][[model_]]</pre>
# #save models estimated by elect() in a external object for faster access in the future
# saveRDS(models, pasteO(pathSaveFolder, "models.rds"))
models <- readRDS(paste0(pathSaveFolder, "models.rds"))</pre>
# inspect created object
lapply(models, names)
$age
[1] "msm" "LE"
$age_bl
[1] "msm" "LE"
$male
[1] "msm" "LE"
$educat
[1] "msm" "LE"
$edu
[1] "msm" "LE"
```

Model results

age

summary

```
model <- models[["age"]]
msm_summary(model$msm)</pre>
```

```
-2loglik = 15059.61
Convergence code = 0
            se Wald ChiSq Pr>ChiSq
         р
gbase -2.12 0.07
                   865.97
                              0.00
qbase -4.13 0.17
                    558.14
                              0.00
                              0.00
qbase -0.29 0.08
                    12.28
qbase -2.19 0.15
                  199.54
                              0.00
qbase -3.06 0.35
                   78.06
                              0.00
                134.20
160.84
qbase -2.12 0.18
                              0.00
qcov 0.08 0.01
                              0.00
                  31.93
                              0.00
qcov 0.08 0.01
qcov -0.02 0.01
                    7.44
                            0.01
```

```
qcov 0.05 0.01
                     23.01
                              0.00
                     6.34
                              0.01
qcov 0.06 0.02
qcov 0.07 0.01
                     33.05
                              0.00
     -2.15 0.09
                    589.75
                              0.00
initp -1.33 0.06
                    433.56
                              0.00
                              0.00
initp -2.85 0.13
                    480.86
```

```
solution
print(model$msm, showEnv= F)
Call:
msm(formula = state ~ age, subject = id, data = ds, qmatrix = Q,
                                                                    ematrix = E, covariates = covariat
Maximum likelihood estimates
Baselines are with covariates set to 0
Transition intensities with hazard ratios for each covariate
                  Baseline
State 1 - State 1 -0.13609 (-0.15443,-0.11992)
State 1 - State 2 0.11996 (0.10416, 0.13816) 1.0814 (1.0684,1.0946)
State 1 - State 4 0.01613 (0.01145, 0.02271) 1.0869 (1.0559,1.1187)
State 2 - State 1 0.74949 (0.63785, 0.88065) 0.9802 (0.9662,0.9944)
State 2 - State 2 -0.90846 (-1.04359,-0.79082)
State 2 - State 3 0.11212 (0.08276, 0.15189) 1.0560 (1.0328,1.0798)
State 2 - State 4 0.04685 (0.02376, 0.09239) 1.0615 (1.0133,1.1119)
State 3 - State 3 -0.11946 (-0.17114,-0.08339)
State 3 - State 4 0.11946 (0.08339, 0.17114) 1.0674 (1.0439,1.0914)
Warning in res[plabs == "pbase"][unique(states[plabs == "p"])] <- 1/(1 + : number of items to replace i
of replacement length
Warning in res[plabs == "pbase"][unique(states[plabs == "p"])] <- 1/(1 + : number of items to replace i
of replacement length
Misclassification probabilities
                      Baseline
Obs State 2 | State 2 0
Obs State 3 | State 2 0.1041 (0.08898,0.1215)
Initial state occupancy probabilities
         Estimate
                         LCL
                                     UCL
```

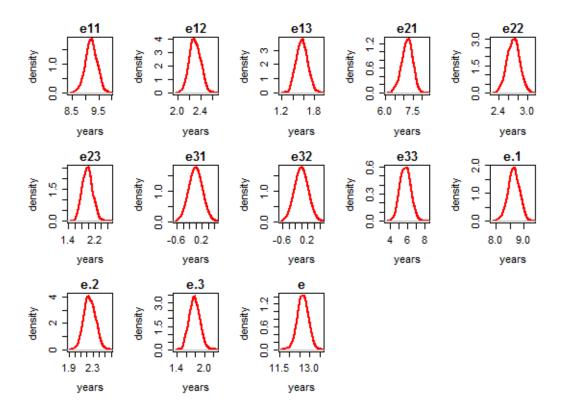
Estimate LCL UCL
State 1 0.75681606 0.73449835 0.77726093
State 2 0.19930412 0.17966161 0.22057570
State 3 0.04387982 0.03430043 0.05598259
State 4 0.00000000 0.00000000 0.00000000

-2 * log-likelihood: 15059.61 [Note, to obtain old print format, use "printold.msm"]

ELECT summary

```
summary.elect(
 model$LE, # life expectancy estimated by elect()
 probs = c(.025, .5, .975), # numeric vector of probabilities for quantiles
 digits=2, # number of decimals places in output
 print = TRUE # print toggle
ELECT summary
Covariates values in the multi-state model:
age
 0
Covariates in the state-distribution model:
Life expectancies: Using simulation with 1000 replications
Point estimates, and mean, SEs, and quantiles from simulation:
     pnt
          mn se 0.025q 0.5q 0.975q
e11 9.26 9.23 0.22 8.78 9.24
                                 9.64
e12 2.33 2.31 0.10 2.12 2.30
                                 2.49
e13 1.58 1.57 0.10 1.38 1.57
                                1.77
e21 7.23 7.20 0.30 6.55 7.21
                                7.73
e22 2.74 2.71 0.13 2.45 2.71
                                 2.95
e23 2.00 1.99 0.15 1.71 1.99
                                 2.30
e31 0.00 0.00 0.00 0.00 0.00
                                 0.00
e32 0.00 0.00 0.00 0.00 0.00
                                 0.00
e33 5.75 5.76 0.62 4.62 5.75
                                 6.98
e.1 8.67 8.64 0.21 8.21 8.65
                                 9.04
e.2 2.27 2.25 0.09 2.07 2.24
                                 2.44
e.3 1.79 1.79 0.11 1.57 1.79 2.01
e 12.73 12.68 0.27 12.16 12.68 13.18
plots
```

```
plot.elect(
  model$LE, # life expectancy estimated by elect()
  kernel = "gaussian", #character string for smoothing kernal ("gaussian",...)
  col = "red", # color of the curve
  lwd = 2, # line width of the curve
  cex.lab = 1 # magnification to be used for axis-labels
)
```



age at baseline

summary

```
model <- models[["age_bl"]]
msm_summary(model$msm)</pre>
```

```
-2loglik = 15026.84
Convergence code = 0
              se Wald ChiSq Pr>ChiSq
qbase -2.02 0.08
                      679.18
                                 0.00
                                 0.00
qbase -4.25 0.20
                      447.08
qbase -0.19 0.09
                        4.94
                                 0.03
                                 0.00
qbase -2.25 0.16
                      196.77
qbase -3.10 0.34
                       81.39
                                 0.00
qbase -2.22 0.19
                      132.69
                                 0.00
       0.04 0.01
                        6.96
                                 0.01
qcov
       0.12 0.03
                       13.74
                                 0.00
qcov
                                 0.00
      -0.07 0.02
                       16.64
qcov
qcov
       0.08 0.02
                       12.07
                                 0.00
       0.11 0.04
                        6.46
                                 0.01
qcov
                       20.14
                                 0.00
qcov
       0.09 0.02
                                 0.00
       0.05 0.01
                       13.09
qcov
qcov
                                 0.23
     -0.04 0.03
                        1.47
       0.06 0.02
                       10.64
                                 0.00
qcov
qcov -0.03 0.02
                        1.49
                                 0.22
qcov -0.07 0.05
                        2.30
                                 0.13
```

```
    qcov
    -0.04
    0.02
    2.65
    0.10

    p
    -2.15
    0.09
    588.84
    0.00

    initp
    -1.33
    0.06
    433.67
    0.00

    initp
    -2.83
    0.13
    487.60
    0.00
```

State 4 0.00000000 0.00000000 0.00000000

[Note, to obtain old print format, use "printold.msm"]

-2 * log-likelihood: 15026.84

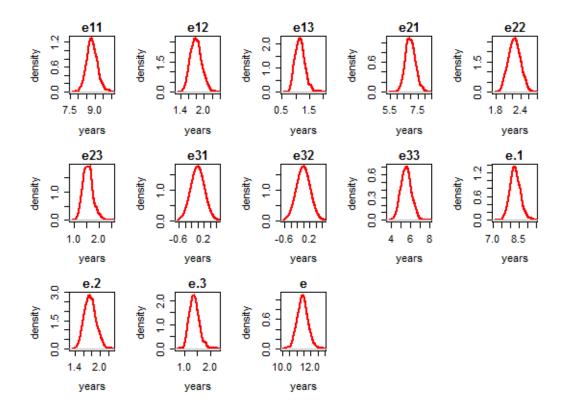
solution

```
print(model$msm, showEnv= F)
Call:
msm(formula = state ~ age, subject = id, data = ds, qmatrix = Q,
                                                                     ematrix = E, covariates = covariate
Maximum likelihood estimates
Baselines are with covariates set to 0
Transition intensities with hazard ratios for each covariate
                 Baseline
                                                                       age_bl
State 1 - State 1 -0.14696 (-0.168443,-0.12821)
State 1 - State 2 0.13272 (0.114020, 0.15449) 1.0361 (1.0092,1.0638) 1.0519 (1.0234,1.081)
State 1 - State 4 0.01423 (0.009598, 0.02111) 1.1269 (1.0579,1.2004) 0.9605 (0.8999,1.025)
State 2 - State 1 0.82371 (0.694275, 0.97728) 0.9317 (0.9006,0.9639) 1.0628 (1.0246,1.102)
State 2 - State 2 -0.97424 (-1.130972,-0.83923)
State 2 - State 3 0.10552 (0.077071, 0.14448) 1.0812 (1.0346,1.1299) 0.9720 (0.9287,1.017)
State 2 - State 4 0.04500 (0.022944, 0.08827) 1.1185 (1.0260,1.2194) 0.9332 (0.8534,1.020)
State 3 - State 3 -0.10897 (-0.158895,-0.07473)
State 3 - State 4 0.10897 (0.074732, 0.15889) 1.0985 (1.0544,1.1445) 0.9652 (0.9249,1.007)
Warning in res[plabs == "pbase"][unique(states[plabs == "p"])] <- 1/(1 + : number of items to replace i
of replacement length
Warning in res[plabs == "pbase"][unique(states[plabs == "p"])] <- 1/(1 + : number of items to replace i
of replacement length
Misclassification probabilities
                      Baseline
Obs State 2 | State 2 0
Obs State 3 | State 2 0.1041 (0.08896,0.1215)
Initial state occupancy probabilities
         Estimate
                         LCI.
State 1 0.75610249 0.73399586 0.77561052
State 2 0.19930454 0.18017069 0.22017770
State 3 0.04459297 0.03486444 0.05662713
```

ELECT summary

```
summary.elect(
 model$LE, # life expectancy estimated by elect()
 probs = c(.025, .5, .975), # numeric vector of probabilities for quantiles
 digits=2, # number of decimals places in output
 print = TRUE # print toggle
ELECT summary
Covariates values in the multi-state model:
  age age_bl
    0 0
Covariates in the state-distribution model:
Life expectancies: Using simulation with 1000 replications
Point estimates, and mean, SEs, and quantiles from simulation:
     pnt
         mn se 0.025q 0.5q 0.975q
e11 8.87 8.83 0.32 8.23 8.81
                                 9.50
e12 1.83 1.80 0.14 1.55 1.79
                                 2.09
e13 1.17 1.16 0.18 0.87 1.15
                                1.56
e21 7.03 6.98 0.35 6.34 6.97
                                7.76
e22 2.29 2.25 0.14 1.99 2.25
                               2.53
e23 1.58 1.57 0.20 1.22 1.56
                                 2.04
e31 0.00 0.00 0.00 0.00 0.00
                                 0.00
e32 0.00 0.00 0.00 0.00 0.00
                                 0.00
e33 5.58 5.58 0.55 4.54 5.58
                                 6.69
e.1 8.32 8.28 0.31 7.71 8.26
                                 8.92
e.2 1.80 1.77 0.13 1.53 1.76
                                 2.04
e.3 1.39 1.39 0.18 1.08 1.38 1.79
e 11.51 11.43 0.37 10.75 11.43 12.18
plots
```

```
plot.elect(
  model$LE, # life expectancy estimated by elect()
  kernel = "gaussian", #character string for smoothing kernal ("gaussian",...)
  col = "red", # color of the curve
  lwd = 2, # line width of the curve
  cex.lab = 1 # magnification to be used for axis-labels
)
```



male

summary

```
model <- models[["male"]]
msm_summary(model$msm)</pre>
```

```
-2loglik = 14977.86
Convergence code = 0
              se Wald ChiSq Pr>ChiSq
qbase -2.10 0.08
                      667.99
                                 0.00
                                 0.00
qbase -4.38 0.21
                      437.79
qbase -0.23 0.09
                        5.94
                                 0.01
                      179.55
                                 0.00
qbase -2.23 0.17
qbase -3.29 0.38
                       74.60
                                 0.00
                      132.89
                                 0.00
qbase -2.47 0.21
       0.04 0.01
                        6.79
                                 0.01
qcov
       0.13 0.03
                       15.41
                                 0.00
qcov
                                 0.00
      -0.07 0.02
                       15.98
qcov
qcov
       0.09 0.02
                       14.44
                                 0.00
       0.09 0.05
                        4.06
                                 0.04
qcov
                                 0.00
qcov
       0.11 0.02
                       27.79
                                 0.00
       0.05 0.01
                       13.23
qcov
                                 0.15
     -0.05 0.03
                        2.12
qcov
       0.06 0.02
                       10.04
                                 0.00
qcov
qcov
     -0.04 0.02
                        2.46
                                 0.12
                                 0.35
qcov -0.05 0.05
                        0.87
```

```
qcov -0.05 0.02
                     5.80
                              0.02
     0.33 0.09
                     13.14
                              0.00
qcov
qcov 0.45 0.23
                     3.77
                              0.05
qcov 0.12 0.11
                      1.30
                              0.25
qcov -0.16 0.16
                      1.08
                              0.30
qcov 0.64 0.28
                     5.13
                              0.02
                              0.00
qcov
      0.43 0.14
                     9.23
     -2.15 0.09
                    589.03
                              0.00
initp -1.33 0.06
                   434.12
                              0.00
initp -2.84 0.13
                    486.43
                              0.00
```

solution

```
print(model$msm, showEnv= F)
Call:
```

Maximum likelihood estimates

Baselines are with covariates set to 0

Baseline

Transition intensities with hazard ratios for each covariate

msm(formula = state ~ age, subject = id, data = ds, qmatrix = Q,

```
State 1 - State 1 -0.13442 (-0.155254,-0.11639)

State 1 - State 2 0.12190 (0.103921, 0.14300) 1.0357 (1.0087,1.0635) 1.0523 (1.0238,1.0817) 1.3930 (1

State 1 - State 4 0.01252 (0.008307, 0.01887) 1.1332 (1.0646,1.2062) 0.9528 (0.8927,1.0169) 1.5623 (0

State 2 - State 1 0.79683 (0.663817, 0.95650) 0.9327 (0.9014,0.9651) 1.0612 (1.0229,1.1009) 1.1290 (0

State 2 - State 2 -0.94185 (-1.104543,-0.80312)

State 2 - State 3 0.10788 (0.077894, 0.14942) 1.0892 (1.0422,1.1383) 0.9642 (0.9212,1.0091) 0.8497 (0

State 2 - State 4 0.03714 (0.017591, 0.07841) 1.0979 (1.0026,1.2024) 0.9558 (0.8692,1.0511) 1.8957 (1

State 3 - State 4 0.08480 (0.055742, 0.12900) 1.1198 (1.0737,1.1679) 0.9489 (0.9093,0.9903) 1.5390 (1
```

ematrix = E, covariates = covariat

male

age_bl

Warning in res[plabs == "pbase"][unique(states[plabs == "p"])] <- 1/(1 + : number of items to replace i of replacement length

Warning in res[plabs == "pbase"][unique(states[plabs == "p"])] <- 1/(1 + : number of items to replace i of replacement length

Misclassification probabilities

Baseline

Obs State 2 | State 2 0

Obs State 3 | State 2 0.1042 (0.08905,0.1216)

Initial state occupancy probabilities

Estimate LCL UCL

State 1 0.75644307 0.7336018 0.77663358

State 2 0.19914669 0.1799655 0.22019658

State 3 0.04441024 0.0350617 0.05654718

State 4 0.00000000 0.0000000 0.00000000

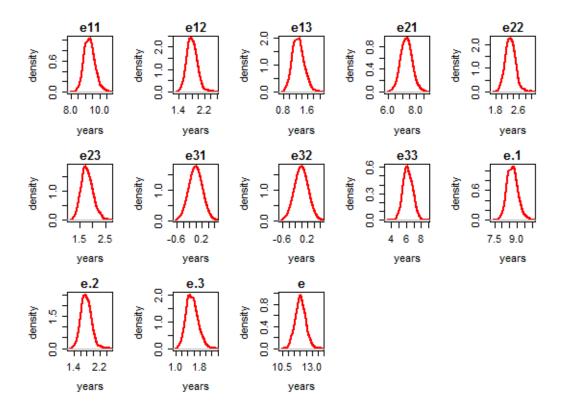
-2 * log-likelihood: 14977.86

[Note, to obtain old print format, use "printold.msm"]

ELECT summary

```
summary.elect(
 model$LE, # life expectancy estimated by elect()
 probs = c(.025, .5, .975), # numeric vector of probabilities for quantiles
 digits=2, # number of decimals places in output
 print = TRUE # print toggle
ELECT summary
Covariates values in the multi-state model:
  age age_bl male
    0
          0
                 0
Covariates in the state-distribution model:
  age
Life expectancies: Using simulation with 1000 replications
Point estimates, and mean, SEs, and quantiles from simulation:
           mn se 0.025q 0.5q 0.975q
     pnt
e11 9.35 9.29 0.37 8.63 9.28 10.09
e12 1.85 1.81 0.16 1.51 1.80
                                2.13
e13 1.31 1.29 0.19
                   0.96 1.28
                                1.68
e21 7.40 7.31 0.40 6.52 7.31
                               8.13
e22 2.34 2.30 0.17 1.98 2.29
                                2.64
e23 1.81 1.80 0.22 1.41 1.78
                                2.28
e31 0.00 0.00 0.00 0.00 0.00
                                0.00
e32 0.00 0.00 0.00 0.00 0.00
                                0.00
e33 6.24 6.26 0.65 5.02 6.22
                                7.54
e.1 8.76 8.70 0.35 8.08 8.69
                                9.47
e.2 1.82 1.78 0.15 1.50 1.77
                                2.08
e.3 1.57 1.55 0.19 1.22 1.54
                               1.96
e 12.15 12.04 0.43 11.20 12.03 12.90
_____
plots
```

```
plot.elect(
  model$LE, # life expectancy estimated by elect()
  kernel = "gaussian", #character string for smoothing kernal ("gaussian",...)
  col = "red", # color of the curve
  lwd = 2, # line width of the curve
  cex.lab = 1 # magnification to be used for axis-labels
)
```



education

summary

```
model <- models[["educat"]]
msm_summary(model$msm)</pre>
```

```
-2loglik = 14965.72
Convergence code = 0
              se Wald ChiSq Pr>ChiSq
qbase -1.87 0.11
                      265.81
                                 0.00
                                 0.00
qbase -4.32 0.32
                      186.83
qbase -0.23 0.11
                        4.07
                                 0.04
                                 0.00
qbase -2.15 0.19
                      125.35
qbase -3.47 0.49
                       49.52
                                 0.00
                      112.26
                                 0.00
qbase -2.38 0.22
       0.04 0.01
                        6.79
                                 0.01
qcov
       0.12 0.03
                       14.88
                                 0.00
qcov
                                 0.00
      -0.07 0.02
                       16.22
qcov
qcov
       0.08 0.02
                       14.06
                                 0.00
       0.10 0.05
                        4.23
                                 0.04
qcov
                                 0.00
qcov
       0.11 0.02
                       28.29
                                 0.00
       0.05 0.01
                       12.34
qcov
                                 0.17
     -0.05 0.03
                        1.87
qcov
       0.06 0.02
                                 0.00
qcov
                       10.14
qcov
     -0.03 0.02
                        2.14
                                 0.14
qcov -0.05 0.05
                                 0.30
                        1.08
```

```
qcov -0.05 0.02
                      5.71
                               0.02
                     12.92
                               0.00
      0.33 0.09
qcov
qcov
      0.49 0.23
                      4.41
                               0.04
qcov 0.12 0.11
                      1.30
                               0.25
qcov -0.14 0.16
                      0.81
                               0.37
qcov 0.57 0.30
                      3.63
                               0.06
                               0.00
qcov
     0.44 0.14
                      9.66
qcov -0.25 0.09
                      7.31
                               0.01
qcov -0.09 0.27
                      0.11
                               0.74
qcov
     0.02 0.11
                      0.03
                               0.87
qcov -0.10 0.14
                      0.56
                               0.45
                      0.33
                               0.56
qcov
     0.25 0.44
qcov -0.15 0.12
                      1.39
                               0.24
     -2.15 0.09
                    588.76
                               0.00
                               0.00
initp -1.33 0.06
                    433.80
initp -2.83 0.13
                    486.75
                               0.00
```

solution

```
print(model$msm, showEnv= F)
```

Call:

msm(formula = state ~ age, subject = id, data = ds, qmatrix = Q, ematrix = E, covariates = covariat

Maximum likelihood estimates

Baselines are with covariates set to 0

Transition intensities with hazard ratios for each covariate

```
Baseline
                                                                      age_bl
                                                                                             male
State 1 - State 1 -0.16778 (-0.205801,-0.13679)
State 1 - State 2 0.15445 (0.123388, 0.19334) 1.0360 (1.0088,1.0640) 1.0510 (1.0222,1.0805) 1.3914 (1
State 1 - State 4 0.01333 (0.007177, 0.02476) 1.1327 (1.0632,1.2067) 0.9546 (0.8931,1.0204) 1.6254 (1
State 2 - State 1 0.79401 (0.634537, 0.99357) 0.9317 (0.9001,0.9643) 1.0617 (1.0233,1.1015) 1.1304 (0
State 2 - State 2 -0.94182 (-1.146462,-0.77370)
State 2 - State 3 0.11662 (0.080061, 0.16989) 1.0880 (1.0411,1.1371) 0.9662 (0.9228,1.0117) 0.8667 (0
State 2 - State 4 0.03118 (0.011869, 0.08192) 1.1008 (1.0046,1.2063) 0.9492 (0.8603,1.0472) 1.7758 (0
State 3 - State 3 -0.09296 (-0.144258,-0.05990)
State 3 - State 4 0.09296 (0.059902, 0.14426) 1.1218 (1.0753,1.1703) 0.9490 (0.9091,0.9906) 1.5593 (1
State 1 - State 1
State 1 - State 2 0.7751 (0.6445,0.9323)
State 1 - State 4 0.9152 (0.5427,1.5433)
State 2 - State 1 1.0175 (0.8271,1.2518)
```

State 2 - State 2

State 2 - State 3 0.9007 (0.6848,1.1847)

State 2 - State 4 1.2886 (0.5450,3.0466)

State 3 - State 3

State 3 - State 4 0.8635 (0.6765,1.1022)

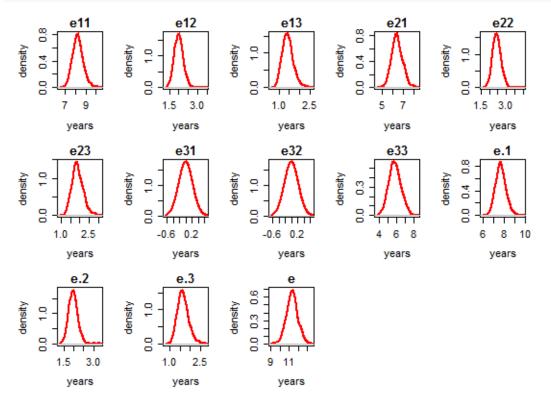
Warning in res[plabs == "pbase"][unique(states[plabs == "p"])] <- 1/(1 + : number of items to replace i of replacement length

```
Warning in res[plabs == "pbase"][unique(states[plabs == "p"])] <- 1/(1 + : number of items to replace i
of replacement length
Misclassification probabilities
                    Baseline
Obs State 2 | State 2 0
Obs State 3 | State 2 0.1042 (0.08904,0.1216)
Initial state occupancy probabilities
         Estimate
                        LCL
                                  UCL
State 1 0.75627049 0.73396526 0.77637390
State 2 0.19927786 0.17984784 0.22039809
State 3 0.04445165 0.03500507 0.05651733
State 4 0.00000000 0.00000000 0.00000000
-2 * log-likelihood: 14965.72
[Note, to obtain old print format, use "printold.msm"]
ELECT summary
summary.elect(
 model$LE, # life expectancy estimated by elect()
 probs = c(.025, .5, .975), # numeric vector of probabilities for quantiles
 digits=2, # number of decimals places in output
 print = TRUE # print toggle
_____
ELECT summary
_____
Covariates values in the multi-state model:
  age age_bl male educat
    0
         0
              0
Covariates in the state-distribution model:
Life expectancies: Using simulation with 1000 replications
Point estimates, and mean, SEs, and quantiles from simulation:
           mn se 0.025q 0.5q 0.975q
     pnt
                    7.31 8.22
e11 8.33 8.24 0.51
                                 9.33
e12 2.07 2.02 0.23 1.58 2.01
                                 2.49
e13 1.45 1.43 0.26 1.00 1.41
                                 1.98
e21 6.56 6.46 0.50 5.56 6.43
                                 7.46
e22 2.51 2.45 0.24
                    1.99 2.44
                                 2.94
e23 1.92 1.90 0.30
                    1.37 1.87
                                 2.55
e31 0.00 0.00 0.00 0.00 0.00
                                0.00
e32 0.00 0.00 0.00 0.00 0.00
                                 0.00
e33 5.84 5.84 0.69 4.57 5.83
                                 7.25
e.1 7.80 7.72 0.48 6.85 7.69
                                 8.75
e.2 2.03 1.97 0.22 1.56 1.97
                                 2.43
e.3 1.68 1.66 0.26 1.20 1.64
                                 2.23
```

```
11.51 11.36 0.60 10.22 11.36 12.55
```

plots

```
plot.elect(
  model$LE, # life expectancy estimated by elect()
  kernel = "gaussian", #character string for smoothing kernal ("gaussian",...)
  col = "red", # color of the curve
  lwd = 2, # line width of the curve
  cex.lab = 1 # magnification to be used for axis-labels
)
```



Session Info

```
sessionInfo()
```

R version 3.3.1 (2016-06-21)

Platform: x86_64-w64-mingw32/x64 (64-bit) Running under: Windows 10 x64 (build 14393)

locale:

[4] LC_NUMERIC=C

[1] LC_COLLATE=English_United States.1252 LC_CTYPE=English_United States.1252 LC_TIME=English_United States.1252

LC_MONETARY=English_U

attached base packages:

[1] stats graphics grDevices utils datasets methods base

other attached packages:

[1] msm_1.6.1 magrittr_1.5 nnet_7.3-12 knitr_1.14

loaded via a namespace (and not attached):

[1]	Rcpp_0.12.6	formatR_1.4	nloptr_1.0.4	plyr_1.8.4	tools_3.3.1	dig
[7]	lme4_1.1-12	evaluate_0.9	tibble_1.2	gtable_0.2.0	nlme_3.1-128	lat
[13]	mgcv_1.8-14	Matrix_1.2-7.1	DBI_0.5	yaml_2.1.13	parallel_3.3.1	Spa
[19]	mvtnorm_1.0-5	expm_0.999-0	dplyr_0.5.0	stringr_1.1.0	MatrixModels_0.4-1	gri
[25]	R6_2.1.3	survival_2.39-5	rmarkdown_1.0	minqa_1.2.4	ggplot2_2.1.0	car
[31]	scales_0.4.0	htmltools_0.3.5	splines_3.3.1	MASS_7.3-45	assertthat_0.1	pbk
[37]	testit_0.5	colorspace_1.2-6	quantreg_5.26	stringi_1.1.1	lazyeval_0.2.0	mun