

Survival Analysis - Sick-Sicker model

The DARTH workgroup

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Please cite our publications when using this code:

- Jalal H, Pechlivanoglou P, Krijkamp E, Alarid-Escudero F, Enns E, Hunink MG. An Overview of R in Health Decision Sciences. *Med Decis Making*. 2017; 37(3): 735-746. <https://journals.sagepub.com/doi/abs/10.1177/0272989X16686559>
- Krijkamp EM, Alarid-Escudero F, Enns EA, Jalal HJ, Hunink MGM, Pechlivanoglou P. Microsimulation modeling for health decision sciences using R: A tutorial. *Med Decis Making*. 2018;38(3):400-22. <https://journals.sagepub.com/doi/abs/10.1177/0272989X18754513>
- Krijkamp EM, Alarid-Escudero F, Enns E, Pechlivanoglou P, Hunink MM, Jalal H. A Multidimensional Array Representation of State-Transition Model Dynamics. *Med Decis Mak*. 2020;40(2):242-248. doi:10.1177/0272989X19893973

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Change `eval` to `TRUE` if you want to knit this document.

01 Load packages

```
if (!require('pacman')) install.packages('pacman'); library(pacman) # use this package to conveniently
# load (install if required) packages from CRAN
p_load("here", "dplyr", "devtools", "gems", "flexsurv", "survHE", "ggplot2", "msm", "igraph", "mstate",
# load (install if required) packages from GitHub
# install_github("DARTH-git/dampack", force = TRUE) Uncomment if there is a newer version
# install_github("DARTH-git/darthtools", force = TRUE) Uncomment if there is a newer version
p_load_gh("DARTH-git/dampack", "DARTH-git/darthtools")
```

02 Load functions

```
# No function needed
```

03 Input model parameters

```
set.seed(1) # set the seed
v_names_states <- c("H", "S1", "S2", "D") # the model states names

# Model structure
n_t <- 30 # time horizon, 30 cycles
c_l <- 1
n_i <- 100000 # number of simulated individuals
n_s <- length(v_names_states) # the number of health states
d_r <- 0.03 # discount rate of 3% per cycle
v_dw <- 1 / ((1 + d_r) ^ (0:n_t)) # discount weight
v_names_str <- c("no treatment", "treatment") # strategy names

# Event probabilities (per cycle)
# Annual transition probabilities
p_HS1 <- 0.15 # probability of becoming sick when healthy

# Annual probabilities of death
# load age dependent probability
# or use "../data/" if you have a datafolder
p_mort <- read.csv("mortProb_age.csv")
# load age distribution
dist_Age <- read.csv("MyPopulation-AgeDistribution.csv")

# Cost inputs
c_H <- 2000 # cost of one cycle in the healthy state
c_S1 <- 4000 # cost of one cycle in the sick state
c_S2 <- 15000 # cost of one cycle in the sicker state
c_D <- 0 # cost of one cycle in the dead state
c_TrT <- 12000 # cost of treatment (per cycle)

# Utility inputs
```

```

u_H      <- 1      # utility when healthy
u_S1     <- 0.75   # utility when sick
u_S2     <- 0.5    # utility when sicker
u_D      <- 0      # utility when dead
u_Trt    <- 0.95   # utility when sick(er) and being treated

times    <- seq(0, n_t, c_1) # the cycles in years

```

04 Sample individual level characteristicS

04.1 Static characteristics

```

set.seed(2019) # set the seed for the simulation of individual characteristics
v_x          <- runif(n_i, min = 0.95, max = 1.05) # treatment effect modifier at baseline

# sample from age distribution an initial age for every individual
v_age0 <- sample(x = dist_Age$age, prob = dist_Age$prop, size = n_i, replace = TRUE)

# store the information at baseline into a data frame.
df_X      <- data.frame(ID = 1:n_i, x = v_x, Age = v_age0)

```

Survival analysis component

```

# load the Sicker data
data_long <- read.csv("data_long_Sicker.csv", row.names = 1)
head(data_long)

# Multi-state models can be fitted independently for each transition. This is more flexible!

# Create subsets for each transition
data_S1H <-
data_S1S2 <-
data_S1D <-
data_S2D <-

# fit independent models for each transition and pick the one that fits best
#your turn
fit_S1H <-
fit_S1S2 <-
fit_S1D <-
fit_S2D <-

best.fit_S1H <-
best.fit_S1S2 <-
best.fit_S1D <-
best.fit_S2D <-

# Extract transition probabilities from the best fitting models
p_S1H <-
p_S1S2 <-

```

```
p_S1D <-  
p_S2D <-
```

04.2 Dynamic characteristics

```
# Specify the initial health state of the individuals  
# everyone begins in the healthy state (in this example)  
v_M_init <- rep("H", n_i) # a vector with the initial health state for all individuals  
v_Ts1_init <- rep(0, n_i) # a vector with the time of being sick at the start of the model  
v_Ts2_init <- rep(0, n_i) # a vector with the time of being sick at the start of the model
```

05 Define Simulation Functions

05.1 Probability function

The function that updates the transition probabilities of every cycle is shown below.

```
# your turn
```

05.2 Cost function

The Costs function estimates the costs at every cycle.

```
# your turn
```

05.3 Health outcome function

The Effs function to update the utilities at every cycle.

```
# your turn
```

06 Run Microsimulation

```
# your turn
```

07 Visualize results

```
# your turn
```

08 Cost Effectiveness Analysis

your turn