Microsimulation Sick-Sicker model with time dependency with PSA

Includes individual characteristics: age, age dependent mortality probabilities, individual treatment effect modifyer, time dependency for the sick (S1) state, increasing change of death in the first 6 year of sickness (tunnel)

The DARTH workgroup

Developed by the Decision Analysis in R for Technologies in Health (DARTH) workgroup:

Fernando Alarid-Escudero, PhD (1)

Eva A. Enns, MS, PhD (2)

M.G. Myriam Hunink, MD, PhD (3,4)

Hawre J. Jalal, MD, PhD (5)

Eline M. Krijkamp, MSc (3)

Petros Pechlivanoglou, PhD (6,7)

Alan Yang, MSc (7)

In collaboration of:

- 1. Drug Policy Program, Center for Research and Teaching in Economics (CIDE) CONACyT, Aguas-calientes, Mexico
- 2. University of Minnesota School of Public Health, Minneapolis, MN, USA
- 3. Erasmus MC, Rotterdam, The Netherlands
- 4. Harvard T.H. Chan School of Public Health, Boston, USA
- 5. University of Pittsburgh Graduate School of Public Health, Pittsburgh, PA, USA
- 6. University of Toronto, Toronto ON, Canada
- 7. The Hospital for Sick Children, Toronto ON, Canada

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- 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 Making. 2020 Online first. https://doi.org/10.1177/0272989X19893973

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

```
rm(list = ls())  # clear memory (removes all the variables from the workspace)
```

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", "scales", "ellipse", "ggplot2", "lazyeval", "igraph", "re
# 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 functions needed
```

03 Input model parameters

```
set.seed(1) # set the seed
# Model structure
n_t <- 30
                                  # time horizon, 30 cycles
n_i <- 100000
                                 # number of simulated individuals
v_n <- c("H", "S1", "S2", "D") # the model states names
n_s <- length(v_n)
                                # the number of health states
                                  # discount rate of 3% per cycle
d r
     <- 0.03
v_dwe \leftarrow v_dwc \leftarrow 1 / ((1 + d_r) ^ (0:n_t)) # discount weight
v_names_str <- c("no treatment", "treatment") # strategy names</pre>
                                # number of strategies
n_str <- length(v_names_str)</pre>
### Event probabilities (per cycle)
# Annual transition probabilities
p_HS1 <- 0.15
                                  # probability of becoming sick when healthy
p_S1H <- 0.5
                                  # probability of recovering to healthy when sick
p_S1S2 <- 0.105
                                  # probability of becoming sicker when sick
# Annual probabilities of death
# load age dependent probability
p_mort <- read.csv("mortProb_age.csv")</pre>
# load age distribution
dist_Age <- read.csv("MyPopulation-AgeDistribution.csv")</pre>
# probability to die in S1 by cycle
```

```
\leftarrow c(0.0149, 0.018, 0.021, 0.026, 0.031, rep(0.037, n_t - 5))
p_S1D
p_S2D
         <- 0.048
                             # probability to die in S2
# 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
        <- 0
                            # cost of one cycle in the dead state
c D
        <- 12000
                            # cost of treatment (per cycle)
c_Trt
# Utility inputs
u_H
       <- 1
                             # utility when healthy
        <- 0.75
                            # utility when sick
u S1
                             # utility when sicker
u_S2
       <- 0.5
\mathtt{u}_{\mathtt{D}}
       <- 0
                             # utility when dead
u_Trt <- 0.95
                             # utility when sick(er) and being treated
```

04 Sample individual level characteristics

04.1 Static characteristics

```
# your turn
v_x <- runif(n_i, min = 0.95, max = 1.05) # treatment effect modifier at baseline</pre>
```

04.2 Dynamic characteristics

```
# your turn
```

04.3 Create a dataframe with the individual characteristics

```
df_X <- #data.frame( # ADD ALL CHARACTERISTICS )</pre>
```

05 Define Simulation Functions

There is no need to make two functions for each strategy. We recomment to make one Probs(), one Cost() and one Effs() function and have an function argument Trt = FALSE which you "switch" on and off of the strategy of interest.

Please see the example below:

```
cost_stay <- function (days = 0, Trt = FALSE) {
  # days: days an individual is staying in a care facility
  # Trt: is the individual treated? (default is FALSE)</pre>
```

```
c_stay_day <- 100  # the price to stay a day at the care facility
c_Trt <- 3000  # the price of treatment, total price for drug. Drug requires one dose

cost <- c_stay_day * days + Trt * c_Trt

return(cost)  # return the pric e
}

cost_stay_noTrt <- cost_stay(days = 10, Trt = FALSE) # run the function for the no treatment strategy
cost_stay_Trt  <- cost_stay(days = 10, Trt = TRUE) # run the function for the treatment strategy

cost_stay_noTrt
cost_stay_Trt</pre>
```

05.1 Probability function

The function that updates the transition probabilities of every cycle is shown below. Please make sure you correctly incorporate the time dependency

```
# your turn
# In this function you have to incorporate age specific mortaility and incorporate the change in probab
```

05.2 Cost function

The Costs function estimates the costs at every cycle.

```
# your turn
# Make sure you incorporate the cost of the treatment in the treatment strategy
```

05.3 Health outcome function

The Effs function to update the utilities at every cycle.

```
# your turn
# Make sure you incorporate the treatment effect modifier
```

06 Run Microsimulation

You have to run the function twice. Once for the treatment strategy and once of the no-treatment strategy

```
# your turn
```

07 Visualize results

your turn

08 Cost Effectiveness Analysis

your turn