

HTA2

```
#install.packages("shiny")
library(shiny)

#ui

ui <- navbarPage(      # creates empty page

  # title of app
  "Markov Model_HTA working group",
  tabPanel("severity distribution",
    numericInput(inputId = "SI_d_severity_MA",
      label = "Severity distribution_MA",
      value = 0.25,
      min = 0,
      max = 1),

    numericInput(inputId = "SI_d_severity_MB",
      label = "Severity distribution_MB",
      value = 0.5,
      min = 0,
      max = 1),

    numericInput(inputId = "SI_d_severity_S",
      label = "Severity distribution_S",
      value = 0.25,
      min = 0,
      max = 1)    ),

  # layout is a sidebar-layout
  # open sidebar panel

  # input type numeric

  tabPanel("country" ,
    radioButtons(inputId ="radio", label = "Country",
      choices = list("Base case" = 1, "Netherlands" = 2, "France"=3),selected = 1)),

  tabPanel("Price of medicine",
    numericInput(inputId = "SI_c_a",
      label = "Price of study medicine A",
      value = 2,
      min = 0,
      max = 20),
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numericInput(inputId = "SI_c_b",
             label = "Price of study medicine B",
             value = 0.5,
             min = 0,
             max = 10)),

tabPanel("maintain care" ,
        fluidRow(
column(2,
    numericInput(inputId = "SI_ru_MA_gp",
                 label = "number of GP visit for mild patients per year",
                 value = 3,
                 min = 0,
                 max = 10),

    numericInput(inputId = "SI_ru_MA_spiro",
                 label = "number of spirometry for mild patients per year",
                 value = 3,
                 min = 0,
                 max = 10),

    numericInput(inputId = "SI_ru_MA_beta",
                 label = "number of beta adrenergics for mild patients per year",
                 value = 100,
                 min = 50,
                 max = 150),

    numericInput(inputId = "SI_ru_MA_theoph",
                 label = "number of theophylline for mild patients per year ",
                 value = 365,
                 min = 100,
                 max = 800),

    numericInput(inputId = "SI_ru_MA_steroid",
                 label = "number of steroid for mild patients per year ",
                 value = 365,
                 min = 100,
                 max = 800)

),

column(2,
    numericInput(inputId = "SI_uc_outgp",
                 label = "unit cost of GP visit",
                 value = 25,
                 min = 10,
                 max = 80),

    numericInput(inputId = "SI_uc_spiro",
                 label = "unit cost of spirometry",
                 value = 20,
                 min = 5,
                 max = 60),

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numericInput(inputId = "SI_uc_beta",
             label = "unit cost of beta adrenergics",
             value = 0.6,
             min = 0.1,
             max = 6),

numericInput(inputId = "SI_uc_theoph",
             label = "unit cost of theophylline ",
             value = 0.5,
             min = 0.1,
             max = 5),

numericInput(inputId = "SI_uc_steroid",
             label = "unit cost of steroid",
             value = 1,
             min = 0.1,
             max = 10),

numericInput(inputId = "SI_uc_outspecialist",
             label = "unit cost of specialist visit",
             value = 60,
             min = 10,
             max = 120)
),

column(2,
      numericInput(inputId = "SI_ru_MB_gp",
                   label = "number of GP visit for moderate patients per year",
                   value = 3,
                   min = 0,
                   max = 10),

      numericInput(inputId = "SI_ru_MB_spiro",
                   label = "number of spirometry for moderate patients per year",
                   value = 3,
                   min = 0,
                   max = 10),

      numericInput(inputId = "SI_ru_MB_beta",
                   label = "number of beta adrenergics for moderate patients per year",
                   value = 150,
                   min = 200,
                   max = 100),

      numericInput(inputId = "SI_ru_MB_theoph",
                   label = "number of theophylline for moderate patients per year ",
                   value = 365,
                   min = 100,
                   max = 800),

      numericInput(inputId = "SI_ru_MB_steroid",
                   label = "number of steroid for moderate patients per year ",
                   value = 365,

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        min = 100,
        max = 800)
    ),
column(2,

numericInput(inputId = "SI_ru_S_gp",
             label = "number of GP visit for servere patients per year",
             value = 4,
             min = 0,
             max = 10),

numericInput(inputId = "SI_ru_S_spiro",
             label = "number of spirometry for servere patients per year",
             value = 4,
             min = 0,
             max = 10),

numericInput(inputId = "SI_ru_S_beta",
             label = "number of beta adrenergics for servere patients per year",
             value = 150,
             min = 200,
             max = 100),

numericInput(inputId = "SI_ru_S_theoph",
             label = "number of theophyline for servere patients per year ",
             value = 365,
             min = 100,
             max = 800),

numericInput(inputId = "SI_ru_S_steroid",
             label = "number of steroid for servere patients per year ",
             value = 365,
             min = 100,
             max = 800)
    ),
column(2,
    numericInput(inputId = "SI_ru_E_h_icu",
                 label = "number of ICU stay for exacerbation ",
                 value = 0,
                 min = 0,
                 max = 5),

    numericInput(inputId = "SI_ru_E_h_noicu",
                 label = "number of non ICU stay for exacerbation ",
                 value = 6,
                 min = 0,
                 max = 10),

    numericInput(inputId = "SI_ru_E_h_er",
                 label = "number of ER visit for exacerbation ",

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        value = 1,
        min = 0,
        max = 5)
    ),
column(2,
    numericInput(inputId = "SI_ru_SE_h_icu",
        label = "number of ICU stay for severe exacerbation ",
        value = 3,
        min = 0,
        max = 7),

    numericInput(inputId = "SI_ru_SE_h_noicu",
        label = "number of non ICU stay for severe exacerbation ",
        value = 8,
        min = 2,
        max = 20),

    numericInput(inputId = "SI_ru_SE_h_er",
        label = "number of ER visit for severe exacerbation ",
        value = 1,
        min = 0,
        max = 5)
    )),
tabPanel("Treatment for exacerbation" ,
    fluidRow(
column(4,
    numericInput(inputId = "SI_uc_h_icu",
        label = "unit cost of ICU stay ",
        value = 1100,
        min = 500,
        max = 2500),

    numericInput(inputId = "SI_uc_h_noicu",
        label = "unit cost of non ICU stay ",
        value = 250,
        min = 120,
        max = 600),

    numericInput(inputId = "SI_uc_h_er",
        label = "unit cost of ER visit ",
        value = 90,
        min = 30,
        max = 200),

    numericInput(inputId = "SI_ru_E_o_pulmon",
        label = "number of specialist visit for exacerbation ",
        value = 1,
        min = 0,
        max = 3),

    numericInput(inputId = "SI_ru_E_o_gp",

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        label = "number of gp visit for exacerbation ",
        value = 1.5,
        min = 0,
        max = 5),

numericInput(inputId = "SI_ru_E_o_anti",
            label = "number of antibiotic use for exacerbation ",
            value = 10,
            min = 0,
            max = 30),

numericInput(inputId = "SI_ru_E_o_syssteroid",
            label = "number of systematic steroid use for exacerbation ",
            value = 15,
            min = 5,
            max = 50),

numericInput(inputId = "SI_ru_E_o_inhsteroid",
            label = "number of inhale steroid use for exacerbation ",
            value = 40,
            min = 5,
            max = 100),

numericInput(inputId = "SI_ru_E_o_theoph",
            label = "number of theophylline use for exacerbation ",
            value = 40,
            min = 5,
            max = 200)
),

column(4,
    numericInput(inputId = "SI_ru_SE_o_pulmon",
                label = "number of specialist visit for servere exacerbation ",
                value = 1.7,
                min = 0,
                max = 5),

    numericInput(inputId = "SI_ru_SE_o_gp",
                label = "number of gp visit for servere exacerbation ",
                value = 1.6,
                min = 0,
                max = 6),

    numericInput(inputId = "SI_ru_SE_o_anti",
                label = "number of antibiotic use for servere exacerbation ",
                value = 16,
                min = 0,
                max = 40),

    numericInput(inputId = "SI_ru_SE_o_syssteroid",
                label = "number of systematic steroid use for servere exacerbation ",
                value = 30,
                min = 5,

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        max = 90),

numericInput(inputId = "SI_ru_SE_o_inhsteroid",
             label = "number of inhale steroid use for servere exacerbation ",
             value = 35,
             min = 5,
             max = 120),

numericInput(inputId = "SI_ru_SE_o_theoph",
             label = "number of theophyline use for servere exacerbation ",
             value = 35,
             min = 5,
             max = 200),

numericInput(inputId = "SI_ru_SE_o_oxy",
             label = "number of oxygen use for servere exacerbation ",
             value = 1,
             min = 0,
             max = 5),

numericInput(inputId = "SI_uc_o_pulmon",
             label = "unit cost of specialist visit for (servere) exacerbation",
             value = 70,
             min = 20,
             max = 200)
),

column(4,
      numericInput(inputId = "SI_uc_o_gp",
                   label = "unit cost of gp visit for (servere) exacerbation ",
                   value = 25,
                   min = 10,
                   max = 70),

      numericInput(inputId = "SI_uc_o_anti",
                   label = "unit cost of antibiotics ",
                   value = 5,
                   min = 0.1,
                   max = 20),

      numericInput(inputId = "SI_uc_o_syssteroid",
                   label = "unit cost of systematic steroid ",
                   value = 0.9,
                   min = 0.1,
                   max = 9),

      numericInput(inputId = "SI_uc_o_inhsteroid",
                   label = "unit cost of inhalation steroid ",
                   value = 0.5,
                   min = 0.1,
                   max = 5),

      numericInput(inputId = "SI_uc_o_theoph",

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        label = "unit cost of theophyline ",
        value = 0.3,
        min = 0.1,
        max = 5),

    numericInput(inputId = "SI_uc_o_oxy",
                  label = "unit cost of oxygen ",
                  value = 235,
                  min = 50,
                  max = 600)

  )),
  tabPanel(
    "select results to show" ,
    radioButtons(inputId = "radio1", label = "Result to show",
                  choices = list("patient time in disease status" = 1, "cost of medicine" = 2, "cost of

    tabPanel(actionButton(inputId = "run_model",
                           label = "Run model") ),
    # action button runs model when pressed

    # close sidebarPanel

    # open main panel
    mainPanel(
      tabsetPanel(
        # heading (results table)

        # tableOutput id = icer_table, from server
        tabPanel("Result Table", tableOutput(outputId = "result_table")),
        tabPanel("Transition Probability", fluidRow(tableOutput(outputId = "prob_table_trtA_first"), tableOutput(
        tabPanel("Probability Exacerbation", tableOutput(outputId = "table_P_ex")),
        tabPanel("Probability Severe Exacerbation", tableOutput(outputId = "table_P_ex_s"))

      )

    ) # close mainpanel

  ) # close UI fluidpage

```

```

## Warning: Navigation containers expect a collection of 'bslib::nav()'/
## 'shiny::tabPanel()'s and/or 'bslib::nav_menu()'/ 'shiny::navbarMenu()'s. Consider
## using 'header' or 'footer' if you wish to place content above (or below) every
## panel's contents.

```


Shiny server function —

```
server <- function(input, output){  
  
  # when action button pressed ...  
  datasetinput<- reactive({  
  
    d_MA <- input$SI_d_severity_MA  
    d_MB <- input$SI_d_severity_MB  
    d_S <- input$SI_d_severity_S  
    country <-input$radio  
  
    ppp<- {if (country== 2){ppp<-1.2}  
    else if (country== 3){ppp<-0.8}  
    else {ppp<-1}}  
  
    show_results<- input$radio1  
  
    show_vars<-{if(show_results==1){show_vars<- c("Time_in_Mild","Time_in_Moderate","Time_in_Severe")}  
    else if(show_results==2){show_vars<- c("Cost_Drug")}  
    else if(show_results==3){show_vars<- c("Cost_maintain")}  
    else if(show_results==4){show_vars<- c("Cost_Hospitalization_exacerbation","Cost_Avoid_exacerbation")}  
    else{show_vars<- c("Exacerbation","Severe_exacerbation","Total_Costs","Avoid_exacerbation")}  
  }  
  
    c_a <- input$SI_c_a  
    c_b <- input$SI_c_b  
    ru_MA_gp <- input$SI_ru_MA_gp  
    ru_MA_spiro <- input$SI_ru_MA_spiro  
    ru_MA_infvac <- 1  
    ru_MA_beta <- input$SI_ru_MA_beta  
    ru_MA_theoph <- input$SI_ru_MA_theoph  
    ru_MA_steroid <- input$SI_ru_MA_steroid  
    pp_MA_gp <- 1  
    pp_MA_spiro <-1  
    pp_MA_infvac <-1  
    pp_MA_beta <- 0.5  
    pp_MA_theoph <-0.5  
    pp_MA_steroid <-0.2  
    uc_outgp <- input$SI_uc_outgp  
    uc_spiro <- input$SI_uc_spiro  
    uc_infvac <- 15  
    uc_beta <- input$SI_uc_beta  
    uc_theoph <- input$SI_uc_theoph  
    uc_steroid <- input$SI_uc_steroid  
  
    ru_MB_gp <- input$SI_ru_MB_gp  
    ru_MB_spiro <- input$SI_ru_MB_spiro  
    ru_MB_infvac <- 1  
    ru_MB_beta <- input$SI_ru_MB_beta  
    ru_MB_theoph <- input$SI_ru_MB_theoph  
    ru_MB_steroid <- input$SI_ru_MB_steroid  
    pp_MB_gp <- 1
```

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pp_MB_spiro <-1
pp_MB_infvac <-1
pp_MB_beta <- 0.6
pp_MB_theoph <-0.6
pp_MB_steroid <-0.4
uc_outspecialist <- input$SI_uc_outspecialist

ru_S_gp <- input$SI_ru_S_gp
ru_S_spiro <- input$SI_ru_S_spiro
ru_S_infvac <- 1
ru_S_beta <- input$SI_ru_S_beta
ru_S_theoph <- input$SI_ru_S_theoph
ru_S_steroid <- input$SI_ru_S_steroid
pp_S_gp <- 1
pp_S_spiro <-1
pp_S_infvac <-1
pp_S_beta <- 0.7
pp_S_theoph <-0.7
pp_S_steroid <-0.7

ru_E_h_icu <- input$SI_ru_E_h_icu
ru_E_h_noicu <- input$SI_ru_E_h_noicu
ru_E_h_er <- input$SI_ru_E_h_er
pp_E_h_icu <- 0
pp_E_h_noicu <- 0.1
pp_E_h_er <- 0.05
uc_h_icu <- input$SI_uc_h_icu
uc_h_noicu <- input$SI_uc_h_noicu
uc_h_er <- input$SI_uc_h_er

ru_E_o_pulmon <- input$SI_ru_E_o_pulmon
ru_E_o_gp <- input$SI_ru_E_o_gp
ru_E_o_anti <- input$SI_ru_E_o_anti
ru_E_o_syssteroid <- input$SI_ru_E_o_syssteroid
ru_E_o_inhsteroid <- input$SI_ru_E_o_inhsteroid
ru_E_o_theoph <- input$SI_ru_E_o_theoph
pp_E_o_pulmon <- 0.35
pp_E_o_gp <- 0.4
pp_E_o_anti <- 0.7
pp_E_o_syssteroid <- 0.5
pp_E_o_inhsteroid <- 0.15
pp_E_o_theoph <- 0.1
uc_o_pulmon <- input$SI_uc_o_pulmon
uc_o_gp <- input$SI_uc_o_gp
uc_o_anti <- input$SI_uc_o_anti
uc_o_syssteroid <- input$SI_uc_o_syssteroid
uc_o_inhsteroid <- input$SI_uc_o_inhsteroid
uc_o_theoph <-input$SI_uc_o_theoph

ru_SE_h_icu <- input$SI_ru_SE_h_icu
ru_SE_h_noicu <- input$SI_ru_SE_h_noicu

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ru_SE_h_er <- input$SI_ru_SE_h_er
pp_SE_h_icu <- 0.1
pp_SE_h_noicu <- 0.75
pp_SE_h_er <- 0.25
uc_h_icu <- input$SI_uc_h_icu
uc_h_noicu <- input$SI_uc_h_noicu
uc_h_er <- input$SI_uc_h_er

ru_SE_o_pulmon <- input$SI_ru_SE_o_pulmon
ru_SE_o_gp <- input$SI_ru_SE_o_gp
ru_SE_o_anti <- input$SI_ru_SE_o_anti
ru_SE_o_syssteroid <- input$SI_ru_SE_o_syssteroid
ru_SE_o_inhsteroid <- input$SI_ru_SE_o_inhsteroid
ru_SE_o_theoph <- input$SI_ru_SE_o_theoph
ru_SE_o_oxy <- input$SI_ru_SE_o_oxy
pp_SE_o_pulmon <- 0.5
pp_SE_o_gp <- 0.4
pp_SE_o_anti <- 0.75
pp_SE_o_syssteroid <- 0.8
pp_SE_o_inhsteroid <- 0.2
pp_SE_o_theoph <- 0.3
pp_SE_o_oxy <- 0.2

uc_o_oxy <- input$SI_uc_o_oxy

# 1 month per cycle
n_cycles <- 12 # number of cycles
v_names_cycles <- paste("cycle", 0:n_cycles) # cycle names
v_names_states <- c("MA", "MB", "S") # state names
n_states <- length(v_names_states) # number of health states
# Strategy names
v_names_str <- c("Treatment A",
                "Treatment B")
n_str <- length(v_names_str) # number of strategies

#transition probability in trt A
#from MA
p_a_MA_MB <- c(0.09, rep(0.04, (n_cycles-1)))
p_a_MA_S <- c(rep(0.01, (n_cycles)))
p_a_MA_MA <- c(0.9, rep(0.95, (n_cycles-1)))

#from MB
p_a_MB_MA <- c(0.25, rep(0.025, (n_cycles-1)))
p_a_MB_S <- c(0.05, rep(0.025, (n_cycles-1)))
p_a_MB_MB <- 1-(p_a_MB_MA+
                p_a_MB_S)

#from S
p_a_S_MA <- c( rep(0.01, (n_cycles)))
p_a_S_MB <-c(0.34, rep(0.04, (n_cycles-1)))
p_a_S_S <- 1-(p_a_S_MA+
                p_a_S_MB)

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#transition probability in trt B
p_b_MA_MB <- c(0.24 ,rep(0.09, (n_cycles-1)))
p_b_MA_S <- c(rep(0.01, (n_cycles)))
p_b_MA_MA <- 1-(p_b_MA_MB+
                p_b_MA_S)

#from MB
p_b_MB_MA <- c(0.1,rep(0.01,(n_cycles-1)))
p_b_MB_S <- c(0.05, rep(0.04,(n_cycles-1)))
p_b_MB_MB <- 1-(p_b_MB_MA+
                p_b_MB_S)

#from S
p_b_S_MA <- c(rep(0.01,(n_cycles)))
p_b_S_MB <-c(0.19, rep(0.02,(n_cycles-1)))
p_b_S_S <- 1-(p_b_S_MA+
              p_b_S_MB)

#probability to ex
#treatment a prob. to ex in trt A

p_a_MA <- 0.05
p_a_MB <- 0.08
p_a_S <- 0.11

#treatment b prob. to ex

p_b_MA <- 0.08
p_b_MB <- 0.1
p_b_S <- 0.12

# prob. of ex to be severe in trt a
p_a_MA_s <- 0.1
p_a_MB_s <- 0.15
p_a_S_s <- 0.2

# prob. of ex to be severe in trt b
p_b_MA_s <- 0.15
p_b_MB_s <- 0.20
p_b_S_s <- 0.25

# maintain cost in trtA per cycle
c_E_h <- ((ru_E_h_icu*pp_E_h_icu*uc_h_icu)+(ru_E_h_noicu*pp_E_h_noicu*uc_h_noicu)+(ru_E_h_er*pp_E_h_er*
c_E_o <- ((ru_E_o_pulmon*pp_E_o_pulmon*uc_o_pulmon)+(ru_E_o_gp*pp_E_o_gp*uc_o_gp)+(ru_E_o_anti*pp_E_o_anti*

c_SE_h <-((ru_SE_h_icu*pp_SE_h_icu*uc_h_icu)+(ru_SE_h_noicu*pp_SE_h_noicu*uc_h_noicu)+(ru_SE_h_er*pp_SE_h_er*
c_SE_o <-((ru_SE_o_pulmon*pp_SE_o_pulmon*uc_o_pulmon)+(ru_SE_o_gp*pp_SE_o_gp*uc_o_gp)+(ru_SE_o_anti*pp_SE_o_anti*

c_MA <- ( ((ru_MA_gp*pp_MA_gp*uc_outgp)+(ru_MA_spiro*pp_MA_spiro*uc_spiro)+(ru_MA_infvac*pp_MA_infvac*uc_
c_MB <- (((ru_MB_gp*pp_MB_gp*uc_outspecialist)+(ru_MB_spiro*pp_MB_spiro*uc_spiro)+(ru_MB_infvac*pp_MB_infvac*uc_

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c_S <- ((ru_S_gp*pp_S_gp*uc_outspecialist)+(ru_S_spiro*pp_S_spiro*uc_spiro)+(ru_S_infvac*pp_S_infvac*uc_infvac))

#drug cost

c_drug_trtA <- c_a*30.5*n_cycles
c_drug_trtB <- c_b*30.5*n_cycles

#hospitalized cost for ex_trtA
hc_ex_MA_trtA <- ((p_a_MA*(1-p_a_MA_s))*(c_E_h))
hc_ex_MB_trtA <- ((p_a_MB*(1-p_a_MB_s))*(c_E_h))
hc_ex_S_trtA <- ((p_a_S*(1-p_a_S_s))*(c_E_h))

#other cost for ex_trtA
oc_ex_MA_trtA <- ((p_a_MA*(1-p_a_MA_s))*(c_E_o))
oc_ex_MB_trtA <- ((p_a_MB*(1-p_a_MB_s))*(c_E_o))
oc_ex_S_trtA <- ((p_a_S*(1-p_a_S_s))*(c_E_o))

#hospitalized cost for severe ex_trtA
hc_exs_MA_trtA <- ((p_a_MA*(p_a_MA_s))*(c_SE_h))
hc_exs_MB_trtA <- ((p_a_MB*(p_a_MB_s))*(c_SE_h))
hc_exs_S_trtA <- ((p_a_S*(p_a_S_s))*(c_SE_h))

#other cost for severe ex_trtA
oc_exs_MA_trtA <- ((p_a_MA*(p_a_MA_s))*(c_SE_o))
oc_exs_MB_trtA <- ((p_a_MB*(p_a_MB_s))*(c_SE_o))
oc_exs_S_trtA <- ((p_a_S*(p_a_S_s))*(c_SE_o))

#hospitalized cost for ex_trtB
hc_ex_MA_trtB <- ((p_b_MA*(1-p_b_MA_s))*(c_E_h))
hc_ex_MB_trtB <- ((p_b_MB*(1-p_b_MB_s))*(c_E_h))
hc_ex_S_trtB <- ((p_b_S*(1-p_b_S_s))*(c_E_h))

#other cost for ex_trtB
oc_ex_MA_trtB <- ((p_b_MA*(1-p_b_MA_s))*(c_E_o))
oc_ex_MB_trtB <- ((p_b_MB*(1-p_b_MB_s))*(c_E_o))
oc_ex_S_trtB <- ((p_b_S*(1-p_b_S_s))*(c_E_o))

#hospitalized cost for severe ex_trtB
hc_exs_MA_trtB <- ((p_b_MA*(p_b_MA_s))*(c_SE_h))
hc_exs_MB_trtB <- ((p_b_MB*(p_b_MB_s))*(c_SE_h))
hc_exs_S_trtB <- ((p_b_S*(p_b_S_s))*(c_SE_h))

#other cost for severe ex_trtB
oc_exs_MA_trtB <- ((p_b_MA*(p_b_MA_s))*(c_SE_o))
oc_exs_MB_trtB <- ((p_b_MB*(p_b_MB_s))*(c_SE_o))
oc_exs_S_trtB <- ((p_b_S*(p_b_S_s))*(c_SE_o))

#cost in trtA per cycle
c_totalMA_trtA <- (c_a*30.5)+
  ((p_a_MA*(1-p_a_MA_s))*(c_E_h+c_E_o))+
  ((p_a_MA*(p_a_MA_s))*(c_SE_h+c_SE_o))+
  (c_MA/12)

```

```

c_totalMB_trtA <- (c_a*30.5)+
  ((p_a_MB*(1-p_a_MB_s))*(c_E_h+c_E_o))+
  ((p_a_MB*(p_a_MB_s))*(c_SE_h+c_SE_o))+
  (c_MB/12)
c_totalS_trtA <- (c_a*30.5)+
  ((p_a_S*(1-p_a_S_s))*(c_E_h+c_E_o))+
  ((p_a_S*(p_a_S_s))*(c_SE_h+c_SE_o))+
  (c_S/12)

#cost in trtB per cycle

c_totalMA_trtB <- (c_b*30.5)+
  ((p_b_MA*(1-p_b_MA_s))*(c_E_h+c_E_o))+
  ((p_b_MA*(p_b_MA_s))*(c_SE_h+c_SE_o))+
  (c_MA/12)

c_totalMB_trtB <- (c_b*30.5)+
  ((p_b_MB*(1-p_b_MB_s))*(c_E_h+c_E_o))+
  ((p_b_MB*(p_b_MB_s))*(c_SE_h+c_SE_o))+
  (c_MB/12)

c_totalS_trtB <- (c_b*30.5)+
  ((p_b_S*(1-p_b_S_s))*(c_E_h+c_E_o))+
  ((p_b_S*(p_b_S_s))*(c_SE_h+c_SE_o))+
  (c_S/12)

## 04.1 Cohort trace
## Initial state vector
# All starting healthy
v_s_init <- c("MA" = d_MA, "MB" = d_MB, "S" = d_S)
v_s_init

## Initialize cohort trace for cSTM for all strategies
m_M_trtA <- matrix(0,
  nrow = (n_cycles + 1), ncol = n_states,
  dimnames = list(v_names_cycles, v_names_states))
# Store the initial state vector in the first row of the cohort trace
m_M_trtA[1, ] <- v_s_init
## Initialize cohort traces
m_M_trtB <- m_M_trtA # structure and initial states remain the same

## 04.2 Transition probability array
## Initialize transition probability array
# all transitions to a non-death state are assumed to be conditional on survival
a_P_trtA <- array(0, # Create 3-D array
  dim = c(n_states, n_states, n_cycles),
  dimnames = list(v_names_states, v_names_states,
    v_names_cycles[-length(v_names_cycles)])) # name the dimensions of the

a_P_trtB <- array(0, # Create 3-D array
  dim = c(n_states, n_states, n_cycles),
  dimnames = list(v_names_states, v_names_states,
    v_names_cycles[-length(v_names_cycles)])) # name the dimensions of the

```

```

## fill the matrix with probs
## trt A
# from MA
a_P_trtA["MA", "MA",] <- p_a_MA_MA
a_P_trtA["MA", "MB",] <- p_a_MA_MB
a_P_trtA["MA", "S",] <- p_a_MA_S

# from MB
a_P_trtA["MB", "MA",] <- p_a_MB_MA
a_P_trtA["MB", "MB",] <- p_a_MB_MB
a_P_trtA["MB", "S",] <- p_a_MB_S

# from S
a_P_trtA["S", "MA",] <- p_a_S_MA
a_P_trtA["S", "MB",] <- p_a_S_MB
a_P_trtA["S", "S",] <- p_a_S_S

## trt B
# from MA
a_P_trtB["MA", "MA",] <- p_b_MA_MA
a_P_trtB["MA", "MB",] <- p_b_MA_MB
a_P_trtB["MA", "S",] <- p_b_MA_S

# from MB
a_P_trtB["MB", "MA",] <- p_b_MB_MA
a_P_trtB["MB", "MB",] <- p_b_MB_MB
a_P_trtB["MB", "S",] <- p_b_MB_S

# from S
a_P_trtB["S", "MA",] <- p_b_S_MA
a_P_trtB["S", "MB",] <- p_b_S_MB
a_P_trtB["S", "S",] <- p_b_S_S

# 05 Run Markov model

for (t in 1:n_cycles){ # loop through the number of cycles
  # estimate the cohort trace for cycle t + 1 using the t-th matrix from the probability array

  m_M_trtA[t + 1, ] <- m_M_trtA[t, ] %*% a_P_trtA[, , t]
  m_M_trtB[t + 1, ] <- m_M_trtB[t, ] %*% a_P_trtB[, , t]
}

df_prob_trtA_first<-data.frame(a_P_trtA[, ,1])
df_prob_trtA_after<-data.frame(a_P_trtA[, ,2])
df_prob_trtB_first<-data.frame(a_P_trtB[, ,1])
df_prob_trtB_after<-data.frame(a_P_trtB[, ,2])

df_prob_ex<-data.frame(p_a_MA, p_a_MB, p_a_S)
df_prob_ex_s<- data.frame(p_a_MA_s, p_a_MB_s, p_a_S_s)

```

```

# print the first few lines of the matrix

## 07.1 Costs and exacerbation nr

# per cycle
# calculate expected costs by multiplying cohort trace with the cost vector for the different health st

v_tc_trtA <- m_M_trtA %*% c(c_totalMA_trtA, c_totalMB_trtA, c_totalS_trtA)
v_tc_trtB <- m_M_trtB %*% c(c_totalMA_trtB, c_totalMB_trtB, c_totalS_trtB)

v_hc_ex_trtA <- m_M_trtA %*% c(hc_ex_MA_trtA, hc_ex_MB_trtA, hc_ex_S_trtA)
v_hc_ex_trtB <- m_M_trtB %*% c(hc_ex_MA_trtB, hc_ex_MB_trtB, hc_ex_S_trtB)

v_oc_ex_trtA <- m_M_trtA %*% c(oc_ex_MA_trtA, oc_ex_MB_trtA, oc_ex_S_trtA)
v_oc_ex_trtB <- m_M_trtB %*% c(oc_ex_MA_trtB, oc_ex_MB_trtB, oc_ex_S_trtB)

v_hc_exs_trtA <- m_M_trtA %*% c(hc_exs_MA_trtA, hc_exs_MB_trtA, hc_exs_S_trtA)
v_hc_exs_trtB <- m_M_trtB %*% c(hc_exs_MA_trtB, hc_exs_MB_trtB, hc_exs_S_trtB)

v_oc_exs_trtA <- m_M_trtA %*% c(oc_exs_MA_trtA, oc_exs_MB_trtA, oc_exs_S_trtA)
v_oc_exs_trtB <- m_M_trtB %*% c(oc_exs_MA_trtB, oc_exs_MB_trtB, oc_exs_S_trtB)

v_mc_trtA <- m_M_trtA %*% c(c_MA, c_MB, c_S)
v_mc_trtB <- m_M_trtB %*% c(c_MA, c_MB, c_S)

# per cycle
# calculate expected exacerbation number
v_exs_trtA <- m_M_trtA %*% c(p_a_MA*p_a_MA_s, p_a_MB*p_a_MB_s, p_a_S*p_a_S_s)
v_exs_trtB <- m_M_trtB %*% c(p_b_MA*p_b_MA_s, p_b_MB*p_b_MB_s, p_b_S*p_b_S_s)

v_ex_trtA <- m_M_trtA %*% c(p_a_MA, p_a_MB, p_a_S)
v_ex_trtB <- m_M_trtB %*% c(p_b_MA, p_b_MB, p_b_S)

## total time in disease status
v_a_MA<- m_M_trtA[, "MA"]
v_time_underdis_MA_a <- sum(v_a_MA)

v_a_MB<- m_M_trtA[, "MB"]
v_time_underdis_MB_a <- sum(v_a_MB)

v_a_S<- m_M_trtA[, "S"]
v_time_underdis_S_a <- sum(v_a_S)

v_b_MA<- m_M_trtB[, "MA"]
v_time_underdis_MA_b <- sum(v_b_MA)

v_b_MB<- m_M_trtB[, "MB"]
v_time_underdis_MB_b <- sum(v_b_MB)

v_b_S<- m_M_trtB[, "S"]
v_time_underdis_S_b <- sum(v_b_S)

df_months_in_disease_state_at_12month <-data.frame(

```



```

        Option = c("Treatment A","Treatment B"),

        Time_in_Mild = c(v_time_underdis_MA_a,
                          v_time_underdis_MA_b),
        Time_in_Moderate = c(v_time_underdis_MB_a,
                              v_time_underdis_MB_b),
        Time_in_Severe = c(v_time_underdis_S_a,
                            v_time_underdis_S_b)
    )

## 06.2.1 total exacerbation nr (u)

v_u_trtA <- sum(v_ex_trtA)
v_u_trtB <- sum(v_ex_trtB)

## 06.2.1 total severe exacerbation nr (u)
v_su_trtA <- sum(v_exs_trtA)
v_su_trtB <- sum(v_exs_trtB)

## 06.2.1 total cost (c)

v_c_trtA <- sum(v_tc_trtA)
v_c_trtB <- sum(v_tc_trtB)

#total hospitalization cost ex

v_thc_trtA<- sum(v_hc_ex_trtA)
v_thc_trtB<- sum(v_hc_ex_trtB)

#total other cost ex

v_toc_trtA<- sum(v_oc_ex_trtA)
v_toc_trtB<- sum(v_oc_ex_trtB)

#total hospitalization cost ex s

v_thcs_trtA<- sum(v_hc_exs_trtA)
v_thcs_trtB<- sum(v_hc_exs_trtB)

#total other cost ex s

v_tocs_trtA<- sum(v_oc_exs_trtA)
v_tocs_trtB<- sum(v_oc_exs_trtB)

#total maintain cost

v_tmc_trtA <- sum(v_mc_trtA)
v_tmc_trtB <- sum(v_mc_trtB)

c_ex_avoid <- (v_c_trtA-v_c_trtB)/(v_u_trtB-v_u_trtA)

Time_in_Mild <-c(v_time_underdis_MA_a,v_time_underdis_MA_b)
Time_in_Moderate <- c(v_time_underdis_MB_a,

```

```

      v_time_underdis_MB_b)
Time_in_Severe <- c(v_time_underdis_S_a,
      v_time_underdis_S_b)
Exacerbation <- c(v_u_trtA,v_u_trtB)
Severe_exacerbation <- c(v_su_trtA,v_su_trtB)
Cost_Drug <- c(c_drug_trtA,c_drug_trtB)
Cost_Hospitalization_exacerbation <- c(v_thc_trtA,v_thc_trtB)
Cost_Other_exacerbation <- c(v_toc_trtA,v_toc_trtB)
Cost_Hospitalization_severe_exacerbation <- c(v_thcs_trtA,v_thcs_trtB)
Cost_Other_severe_exacerbation <- c(v_tocs_trtA,v_tocs_trtB)
Cost_maintain <- c(v_tmc_trtA,v_tmc_trtB)
Total_Costs <- c(v_c_trtA,v_c_trtB)
Avoid_Exacerbation <- c(-(v_u_trtA -v_u_trtB),NA)
Inc.Costs = c(v_c_trtA - v_c_trtB,NA)
Addintional_costs_per_exacerbation_avoid = c(c_ex_avoid, NA)

dt_res_all <-data.table(Time_in_Mild,Time_in_Moderate ,Time_in_Severe,Exacerbation,Severe_exacerbation,
      Cost_Drug,Cost_Hospitalization_exacerbation,Cost_Other_exacerbation,
      Cost_Hospitalization_severe_exacerbation,Cost_Other_severe_exacerbation,
      Cost_maintain,Total_Costs,Avoid_Exacerbation,
      Inc.Costs,Addintional_costs_per_exacerbation_avoid)

df_res_table <- dt_res_all[ # create dataframe

      # row.names = c("Treatment A","Treatment B"),
      ,..show_vars
      ]

})

      # renderTable continuously updates table
output$result_table <- renderTable({
  if (input$run_model>0){isolate(datasetinput())}
})

n_cycles <- 12 # number of cycles
v_names_cycles <- paste("cycle", 0:n_cycles) # cycle names
v_names_states <- c("MA", "MB", "S") # state names
n_states <- length(v_names_states) # number of health states
# Strategy names
v_names_str <- c("Treatment A",
      "Treatment B")
n_str <- length(v_names_str) # number of strategies

#transition probability in trt A

```

```

#from MA
p_a_MA_MB <- c(0.09, rep(0.04, (n_cycles-1)))
p_a_MA_S <- c(rep(0.01, (n_cycles)))
p_a_MA_MA <- c(0.9, rep(0.95, (n_cycles-1)))

#from MB
p_a_MB_MA <- c(0.25, rep(0.025, (n_cycles-1)))
p_a_MB_S <- c(0.05, rep(0.025, (n_cycles-1)))
p_a_MB_MB <- 1-(p_a_MB_MA+
                p_a_MB_S)

#from S
p_a_S_MA <- c( rep(0.01, (n_cycles)))
p_a_S_MB <-c(0.34, rep(0.04, (n_cycles-1)))
p_a_S_S <- 1-(p_a_S_MA+
              p_a_S_MB)

#transition probability in trt B
p_b_MA_MB <- c(0.24 ,rep(0.09, (n_cycles-1)))
p_b_MA_S <- c(rep(0.01, (n_cycles)))
p_b_MA_MA <- 1-(p_b_MA_MB+
                p_b_MA_S)

#from MB
p_b_MB_MA <- c(0.1,rep(0.01, (n_cycles-1)))
p_b_MB_S <- c(0.05, rep(0.04, (n_cycles-1)))
p_b_MB_MB <- 1-(p_b_MB_MA+
                p_b_MB_S)

#from S
p_b_S_MA <- c(rep(0.01, (n_cycles)))
p_b_S_MB <-c(0.19, rep(0.02, (n_cycles-1)))
p_b_S_S <- 1-(p_b_S_MA+
              p_b_S_MB)

#probability to ex
#treatment a prob. to ex in trt A
p_a_MA <- 0.05
p_a_MB <- 0.08
p_a_S <- 0.11

#treatment b prob. to ex
p_b_MA <- 0.08
p_b_MB <- 0.1
p_b_S <- 0.12

# prob. of ex to be servere in trt a
p_a_MA_s <- 0.1
p_a_MB_s <- 0.15
p_a_S_s <- 0.2

```

```

# prob. of ex to be servere in trt b
p_b_MA_s <- 0.15
p_b_MB_s <- 0.20
p_b_S_s <- 0.25

## 04.2 Transition probability array
## Initialize transition probability array
# all transitions to a non-death state are assumed to be conditional on survival
a_P_trtA <- array(0, # Create 3-D array
                  dim = c(n_states, n_states, n_cycles),
                  dimnames = list(v_names_states, v_names_states,
                                  v_names_cycles[-length(v_names_cycles)])) # name the dimensions of the

## fill the matrix with probs
## trt A
# from MA
a_P_trtA["MA", "MA",] <- p_a_MA_MA
a_P_trtA["MA", "MB",] <- p_a_MA_MB
a_P_trtA["MA", "S",] <- p_a_MA_S

# from MB
a_P_trtA["MB", "MA",] <- p_a_MB_MA
a_P_trtA["MB", "MB",] <- p_a_MB_MB
a_P_trtA["MB", "S",] <- p_a_MB_S

# from S
a_P_trtA["S", "MA",] <- p_a_S_MA
a_P_trtA["S", "MB",] <- p_a_S_MB
a_P_trtA["S", "S",] <- p_a_S_S

df_prob_trtA_first<-data.frame(a_P_trtA[, ,1])
df_prob_trtA_after<-data.frame(a_P_trtA[, ,2])

a_P_trtB <- array(0, # Create 3-D array
                  dim = c(n_states, n_states, n_cycles),
                  dimnames = list(v_names_states, v_names_states,
                                  v_names_cycles[-length(v_names_cycles)])) # name the dimensions of the

## trt B
# from MA
a_P_trtB["MA", "MA",] <- p_b_MA_MA
a_P_trtB["MA", "MB",] <- p_b_MA_MB
a_P_trtB["MA", "S",] <- p_b_MA_S

# from MB
a_P_trtB["MB", "MA",] <- p_b_MB_MA
a_P_trtB["MB", "MB",] <- p_b_MB_MB
a_P_trtB["MB", "S",] <- p_b_MB_S

# from S
a_P_trtB["S", "MA",] <- p_b_S_MA
a_P_trtB["S", "MB",] <- p_b_S_MB

```

```
a_P_trtB["S", "S",] <- p_b_S_S
```

```
df_prob_trtB_first<-data.frame(a_P_trtB[,1])
```

```
df_prob_trtB_after<-data.frame(a_P_trtB[,2])
```

```
df_prob_ex<-data.frame(p_a_MA, p_a_MB, p_a_S)
```

```
df_prob_ex_s<- data.frame(p_a_MA_s, p_a_MB_s, p_a_S_s)
```

```
output$prob_table_trtA_first <- renderTable(df_prob_trtA_first)
```

```
output$prob_table_trtA_after <- renderTable(df_prob_trtA_after)
```

```
output$prob_table_trtB_first <- renderTable(df_prob_trtB_first)
```

```
output$prob_table_trtB_after <- renderTable(df_prob_trtB_after)
```

```
output$table_P_ex <- renderTable(df_prob_ex)
```

```
output$table_P_ex_s <- renderTable(df_prob_ex_s)
```

```
} # Server end
```

```
shinyApp(ui, server)
```

```
## PhantomJS not found. You can install it with webshot::install_phantomjs(). If it is installed, please
```