# Causal Inference Methods

# Causal inference - BRCA

hiddenICP with BRCA with 1/3 + 1/3 + 1/3

```
# Reset the environment
rm(list = ls ())
library(InvariantCausalPrediction)
library(AER)
library(CancerSubtypes)
library(miRBaseConverter)
library(miRLAB)
##============
# Set environment variables here
directoryPath <- "C:/Users/phavy022/MyDoc/02CausalInferenceMethods/Experiment-Final/InOut/"</pre>
# directoryPath <- "R/CausalCompare/"</pre>
topk_miR <- 30
topk_mR <- 1500
topk <- 15000
k <- 500
##===========
source(paste(directoryPath, "CausalCompare.R", sep=""))
# 1. Get data set
load(paste(directoryPath, "01BRCA/BRCA_matchedData2.RData", sep=""))
# Get significant miRs and mRs from matchedData
1 <- getDatabyMAD(matchedData, topk_miR, topk_mR)</pre>
# 2. Estimate causal effects
nmiR <- length(l$miRs)</pre>
nmR <- length(1$mRs)
standardizedData <- scale(1$d)</pre>
temp = matrix(nrow = nmR, ncol = nmiR)
## define experimental settings
ExpInd <- c(rep(0,nrow(standardizedData)))</pre>
noOfsample <- nrow(1$d)</pre>
ExpInd[1:round(noOfsample/3)] <- 1</pre>
ExpInd[(round(noOfsample/3)+1):round((2*noOfsample)/3)] <- 2</pre>
ExpInd[(round((2*noOfsample)/3)+1):noOfsample] <- 3</pre>
temp_maximinCoefficients = matrix(nrow = nmR, ncol = nmiR)
temp_pvalues = matrix(nrow = nmR, ncol = nmiR)
# Estimate causal effects
for (i in 1:nmR) {
 Y <- standardizedData[,nmiR+i]
 X <- standardizedData[,c(1:nmiR)]</pre>
```

```
icp <- hiddenICP(X, Y, ExpInd, alpha = 0.1, mode = "asymptotic", intercept=FALSE)</pre>
  for (k in 1:nmiR) {
    temp[i,k] <- icp$betahat[k]</pre>
    temp maximinCoefficients[i,k] <- icp$maximinCoefficients[k]</pre>
    temp_pvalues[i,k] <- icp$pvalues[k]</pre>
 }
}
colnames(temp) <- 1$miRs</pre>
row.names(temp) <- 1$mRs</pre>
## Write the result
# write.csv(temp, file = paste(directoryPath, "ICP_result.csv", sep = ""))
write.csv(temp, file = paste(directoryPath, "hiddenICP_BRCA13_result.csv", sep = ""))
# 3. Validate
#colnames(temp) <- convertmiRs(colnames(temp))</pre>
# Get the result
temp <- read.csv(file = paste(directoryPath, "hiddenICP_BRCA13_result.csv", sep = ""), row.names = 1)
temp <- data.matrix(temp)</pre>
nmiR <- ncol(temp)</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
for(topk in c(500, 1000, 1500, 2000)) {
  top = Extopk(temp, topk)
  write.csv(top, file = paste(directoryPath, "hiddenICP_BRCA13_top_", topk, ".csv", sep = ""))
 topConfirmed = Validation(top, fileName)
  write.csv(topConfirmed[[1]], file = paste(directoryPath, "hiddenICP_BRCA13_topConfirmed_",
                                              topk, ".csv", sep = ""))
}
# Validate for each miRNA
# colnames(temp) <- convertmiRs(colnames(temp))</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
for(k in c(50, 100, 150, 200)) {
  topConfirmed = NULL
 top = NULL
  for(i in 1:nmiR) {
    miRiTopk = bRank(temp, i, k, FALSE)
    top <- rbind(top, miRiTopk)</pre>
    topConfirmedi = Validation(miRiTopk, fileName)
    if(nrow(topConfirmedi[[1]]) != 0) {
      topConfirmed <- rbind(topConfirmed, topConfirmedi[[1]])</pre>
    }
  }
  write.csv(top, file = paste(directoryPath, "hiddenICP_BRCA13_top_eachmiR_", k, ".csv",
                                         sep = ""), row.names = FALSE)
  write.csv(topConfirmed, file = paste(directoryPath, "hiddenICP_BRCA13_topConfirmed_eachmiR_",
                                         k, ".csv",
                                         sep = ""), row.names = FALSE)
}
```

##=========

#### hiddenICP with BRCA - PAM50

```
# Reset the environment
rm(list = ls ())
library(InvariantCausalPrediction)
library(AER)
library(CancerSubtypes)
library(miRBaseConverter)
library(miRLAB)
library(genefu)
# Set environment variables here
directoryPath <- "C:/Users/phavy022/MyDoc/02CausalInferenceMethods/Experiment-Final/InOut/"</pre>
# directoryPath <- "R/CausalCompare/"</pre>
dataDir = "bioclassifier_data"
inputFileName = "inputFile.txt"
topk_miR <- 30
topk_mR <- 1500
source(paste(directoryPath, "CausalCompare.R", sep=""))
# 1. Get data set
load(paste(directoryPath, "01BRCA/BRCA matchedData2.RData", sep=""))
\# Get significant miRs and mRs from matchedData
1 <- getDatabyMAD(matchedData, topk_miR, topk_mR)</pre>
# 2. Estimate causal effects
# Classify samples based on cancer subtypes using Pam50
# Get 50 mRNAs from Pam50
data(pam50)
str(pam50)
fiftymRNAs <- pam50$centroids.map$probe.centroids</pre>
# In matchedData$mRNAs, there are not CDCA1, KNTC2, ORC6L which are in Pam50
# However, CDCA1 ~ NUF2, KNTC2 ~ NDC80, ORC6L ~ ORC6
# In matchedData$mRNAs, replace NUF2 by CDCA1, NDC80 by KNTC2, ORC6 by ORC6L
colnames(matchedData$mRNAs)[which(colnames(matchedData$mRNAs) == "NUF2")] <- "CDCA1"
colnames(matchedData$mRNAs)[which(colnames(matchedData$mRNAs) == "NDC80")] <- "KNTC2"</pre>
colnames(matchedData$mRNAs) [which(colnames(matchedData$mRNAs) == "ORC6")] <- "ORC6L"</pre>
# Get data of 50 mRNAs of Pam50 from matchedData
fiftymRNAsData <- matchedData$mRNAs[, fiftymRNAs]</pre>
prepareData(fiftymRNAsData, directoryPath, dataDir, inputFileName)
###
# input variables for the subtype prediction script
###
library(ctc)
library(heatmap.plus)
```

```
paramDir <- paste(directoryPath, "bioclassifier_R", sep = "") # the location of unchanging files
# such as the function library and main program
inputDir<- paste(directoryPath, dataDir, sep = "") # the location of the data matrix,
# and where output will be located
inputFile<- inputFileName # the input data matrix as a tab delimited text file
short<-"outputFile" # short name that will be used for output files
calibrationParameters <- NA #the column of the "mediansPerDataset.txt" file to use for calibration;
#NA will force centering within the test set \mathfrak S -1 will not do any
#adjustment (when adjustment performed by used)
hasClinical <- FALSE #may include tumor size as second row, with 'T' as the gene name,
#and encoded as binary (0 for size <= 2cm or 1 for size > 2cm)
#set this variable to FALSE if tumor size is not available
collapseMethod<-"mean" # can be mean or iqr (probe with max iqr is selected)
# typically, mean is preferred for long oligo and
# iqr is preferred for short oligo platforms
####
# run the assignment algorithm
source(paste(paramDir, "subtypePrediction_functions.R", sep="/"))
source(paste(paramDir, "subtypePrediction distributed.R", sep="/"))
result <- read.table(paste(directoryPath, "/", dataDir, "/", "outputFile_pam50scores.txt", sep = ""))</pre>
result <- result[-1, -1]
###-----
nmiR <- length(l$miRs)</pre>
nmR <- length(1$mRs)
standardizedData <- scale(1$d)</pre>
temp = matrix(nrow = nmR, ncol = nmiR)
## define experimental settings
ExpInd <- c(rep(0,nrow(standardizedData)))</pre>
noOfsample <- nrow(1$d)</pre>
ExpInd[which(result[, 6] == "Basal")] <- 1</pre>
ExpInd[which(result[, 6] == "Her2")] <- 2</pre>
ExpInd[which(result[, 6] == "LumA")] <- 3</pre>
ExpInd[which(result[, 6] == "LumB")] <- 4</pre>
ExpInd[which(result[, 6] == "Normal")] <- 5</pre>
temp_maximinCoefficients = matrix(nrow = nmR, ncol = nmiR)
temp_pvalues = matrix(nrow = nmR, ncol = nmiR)
# Estimate causal effects
for (i in 1:nmR) {
 Y <- standardizedData[,nmiR+i]
 X <- standardizedData[,c(1:nmiR)]</pre>
  icp <- hiddenICP(X, Y, ExpInd, alpha = 0.1, mode = "asymptotic", intercept=FALSE)</pre>
  for (k in 1:nmiR) {
   temp[i,k] <- icp$betahat[k]</pre>
  }
```

```
colnames(temp) <- 1$miRs</pre>
row.names(temp) <- 1$mRs</pre>
## Write the result
# write.csv(temp, file = paste(directoryPath, "ICP_result.csv", sep = ""))
write.csv(temp, file = paste(directoryPath, "hiddenICP_BRCA_pam50_result.csv", sep = ""))
# 3. Validate
#colnames(temp) <- convertmiRs(colnames(temp))</pre>
# Get the result
temp <- read.csv(file = paste(directoryPath, "hiddenICP_BRCA_pam50_result.csv", sep = ""),</pre>
                 row.names = 1)
temp <- data.matrix(temp)</pre>
nmiR <- ncol(temp)</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
for(topk in c(500, 1000, 1500, 2000)) {
  top = Extopk(temp, topk)
  write.csv(top, file = paste(directoryPath, "hiddenICP_BRCA_pam50_top_", topk, ".csv", sep = ""))
 topConfirmed = Validation(top, fileName)
  write.csv(topConfirmed[[1]], file = paste(directoryPath, "hiddenICP_BRCA_pam50_topConfirmed_",
                                              topk, ".csv", sep = ""))
}
##========
# Validate for each miRNA
# colnames(temp) <- convertmiRs(colnames(temp))</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
for(k in c(50, 100, 150, 200)) {
  topConfirmed = NULL
  top = NULL
  for(i in 1:nmiR) {
    miRiTopk = bRank(temp, i, k, FALSE)
    top <- rbind(top, miRiTopk)</pre>
    topConfirmedi = Validation(miRiTopk, fileName)
    if(nrow(topConfirmedi[[1]]) != 0) {
      topConfirmed <- rbind(topConfirmed, topConfirmedi[[1]])</pre>
    }
  }
  write.csv(top, file = paste(directoryPath, "hiddenICP_BRCA_pam50_top_eachmiR_", k, ".csv",
                                        sep = ""), row.names = FALSE)
  write.csv(topConfirmed, file = paste(directoryPath, "hiddenICP_BRCA_pam50_topConfirmed_eachmiR_",
                                        k, ".csv",
                                        sep = ""), row.names = FALSE)
}
```

# pcalg - ida (idaFast) (& pc) - BRCA

```
# Reset the environment
rm(list = ls())
# Load libraries
library(robustbase)
library(pcalg)
library(Rgraphviz)
library(CancerSubtypes)
library(miRBaseConverter)
library(miRLAB)
##=============
# Set environment variables here
directoryPath <- "C:/Users/phavy022/MyDoc/02CausalInferenceMethods/Experiment-Final/InOut/"
# directoryPath <- "R/CausalCompare/"</pre>
# directoryPath <- "D:/UniSA/CausalCompare/"</pre>
# topk_miR <- 30
# topk_mR <- 500
# topk <- 7500
# k <- 250
topk_miR <- 30
topk_mR <- 1500
topk <- 15000
k <- 500
method <- "min" # value can be "min", "max" or "median"
pcmethod <- "original"</pre>
alpha \leftarrow 0.05
# Load functions
source(paste(directoryPath, "CausalCompare.R", sep=""))
# 1. Get data set
#load(paste(directoryPath, "01BRCA/BRCA_matchedData.RData", sep=""))
#matchedData$miRs <- processmiRName(matchedData$miRs)</pre>
#matchedData$mRNAs <- processmRNAName(matchedData$mRNAs)</pre>
\#save(\textit{matchedData}, \ file=\textit{paste}(\textit{directoryPath}, \ "01BRCA/BRCA\_\textit{matchedData2}.RData", \ sep=""))
load(paste(directoryPath, "01BRCA/BRCA_matchedData2.RData", sep=""))
\# Get significant miRs and mRs from matchedData
1 <- getDatabyMAD(matchedData, topk_miR, topk_mR)</pre>
# 2. Estimate causal effects
nmiR <- length(l$miRs)</pre>
nmR <- length(1$mRs)
standardizedData <- scale(1$d)</pre>
suffStat <- list(C = cor(standardizedData), n = nrow(standardizedData))</pre>
pc.fit <- pc(suffStat, indepTest=gaussCItest, p=ncol(standardizedData), alpha = alpha,</pre>
             skel.method=pcmethod)
# Run idaFast
temp = matrix(nrow = nmR, ncol = nmiR)
for(i in 1:nmiR) {
```

```
eff.estF <- idaFast(i, c((nmiR+1):(nmiR + nmR)), cov(standardizedData), pc.fit@graph)
  for(j in 1:nmR) {
    absValue <- abs(eff.estF[j,])</pre>
    if(method == "min" | method == "max") {
      if(method == "min") {
        index <- which(absValue==min(absValue, na.rm = TRUE))</pre>
      } else { # "max"
        index <- which(absValue==max(absValue, na.rm = TRUE))</pre>
      }
      temp[j,i] <- eff.estF[j,index[1]]</pre>
    } else { # "median"
      temp[j,i] <- median(eff.estF[j,])</pre>
 }
}
colnames(temp) <- l$miRs</pre>
row.names(temp) <- 1$mRs</pre>
## Write the result
write.csv(temp, file = paste(directoryPath, "pc_idaFast_BRCA_result.csv", sep = ""))
# 3. Validate
# Validate the results of the topk targets of miRNAs predicted by the method
# colnames(temp) <- convertmiRs(colnames(temp))</pre>
# Get the result
temp <- read.csv(file = paste(directoryPath, "pc_idaFast_BRCA_result.csv", sep = ""), row.names = 1)
temp <- data.matrix(temp)</pre>
nmiR <- ncol(temp)</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
for(topk in c(500, 1000, 1500, 2000)) {
  top = Extopk(temp, topk)
  write.csv(top, file = paste(directoryPath, "pc_idaFast_BRCA_top_", topk, ".csv", sep = ""))
 topConfirmed = Validation(top, fileName)
  write.csv(topConfirmed[[1]], file = paste(directoryPath, "pc_idaFast_BRCA_topConfirmed_",
                                              topk, ".csv", sep = ""))
}
##=========
# Validate for each miRNA
# colnames(temp) <- convertmiRs(colnames(temp))</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
for(k in c(50, 100, 150, 200)) {
  topConfirmed = NULL
 top = NULL
 for(i in 1:nmiR) {
    miRiTopk = bRank(temp, i, k, FALSE)
    top <- rbind(top, miRiTopk)</pre>
    topConfirmedi = Validation(miRiTopk, fileName)
    if(nrow(topConfirmedi[[1]]) != 0) {
      topConfirmed <- rbind(topConfirmed, topConfirmedi[[1]])</pre>
    }
  }
```

## JointIDA\_direct (beta) - BRCA

```
# Reset the environment
rm(list = ls())
# Load libraries
library(robustbase)
library(pcalg)
library(Rgraphviz)
library(CancerSubtypes)
library(miRBaseConverter)
library(miRLAB)
library(ParallelPC)
library(doParallel)
# Set environment variables here
\#\ directory Path\ <-\ "C:/Users/phavy022/MyDoc/02CausalInferenceMethods/Experiment-Final/InOut/"
# directoryPath <- "R/CausalCompare/"</pre>
directoryPath <- "D:/UniSA/CausalCompare/"</pre>
# topk miR <- 30
# topk_mR <- 500
# topk <- 7500
# k <- 250
topk miR <- 30
topk_mR <- 1500
topk <- 15000
k <- 500
method <- "min" # value can be "min", "max" or "median"
# Load functions
source(paste(directoryPath, "CausalCompare.R", sep=""))
source(paste(directoryPath, "jointIDA_direct.R", sep=""))
# 1. Get data set
load(paste(directoryPath, "01BRCA/BRCA_matchedData2.RData", sep=""))
# Get significant miRs and mRs from matchedData
1 <- getDatabyMAD(matchedData, topk_miR, topk_mR)</pre>
# 2. Estimate causal effects
nmiR <- length(l$miRs)</pre>
nmR <- length(1$mRs)
standardizedData <- scale(1$d)</pre>
```

```
result_jointIDA <- jointIDA_direct(standardizedData, cause = 1:nmiR,
                                    effect = (nmiR+1):(nmiR+nmR),
                                    method=method, pcmethod = "parallel",
                                    num.cores = 8, mem.efficient = FALSE,
                                    alpha = 0.01, technique = "RRC")
## Write the result
write.csv(result jointIDA, file = paste(directoryPath, "jointIDA direct BRCA result.csv", sep = ""))
# 3. Validate
# Validate the results of the topk targets of miRNAs predicted
# by the method
# Get the result
result_jointIDA <- read.csv(file = paste(directoryPath, "jointIDA_direct_BRCA_result.csv", sep = ""),
                            row.names = 1)
result_jointIDA <- data.matrix(result_jointIDA)</pre>
nmiR <- ncol(result_jointIDA)</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
# colnames(result_jointIDA) <- convertmiRs(colnames(result_jointIDA))</pre>
for(topk in c(500, 1000, 1500, 2000)) {
  top = Extopk(result_jointIDA, topk)
  write.csv(top, file = paste(directoryPath, "jointIDA_direct_BRCA_top_", topk,".csv", sep = ""))
  topConfirmed = Validation(top, fileName)
  write.csv(topConfirmed[[1]], file = paste(directoryPath, "jointIDA_direct_BRCA_topConfirmed_",
                                             topk, ".csv", sep = ""))
}
##=========
# Validate for each miRNA
# colnames(result_jointIDA) <- convertmiRs(colnames(result_jointIDA))</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
for(k in c(50, 100, 150, 200)) {
  topConfirmed = NULL
  top = NULL
  for(i in 1:nmiR) {
    miRiTopk = bRank(result jointIDA, i, k, TRUE)
    top <- rbind(top, miRiTopk)</pre>
    topConfirmedi = Validation(miRiTopk, fileName)
    if(nrow(topConfirmedi[[1]]) != 0) {
      topConfirmed <- rbind(topConfirmed, topConfirmedi[[1]])</pre>
    }
  }
  write.csv(top, file = paste(directoryPath, "jointIDA_direct_BRCA_top_eachmiR_", k, ".csv",
                                        sep = ""), row.names = FALSE)
  write.csv(topConfirmed, file = paste(directoryPath, "jointIDA_direct_BRCA_topConfirmed_eachmiR_",
                                        k, ".csv",
                                        sep = ""), row.names = FALSE)
```

#### Lasso in miRLAB with BRCA

```
# Reset the environment
rm(list = ls ())
library(miRBaseConverter)
library(miRLAB)
# Set environment variables here
directoryPath <- "C:/Users/phavy022/MyDoc/02CausalInferenceMethods/Experiment-Final/InOut/"</pre>
# directoryPath <- "R/CausalCompare/"</pre>
topk_miR <- 30
topk mR <- 1500
topk <- 15000
k <- 500
##============
source(paste(directoryPath, "CausalCompare.R", sep=""))
# 1. Get data set
load(paste(directoryPath, "01BRCA/BRCA_matchedData2.RData", sep=""))
# Get significant miRs and mRs from matchedData
1 <- getDatabyMAD(matchedData, topk_miR, topk_mR)</pre>
#l$miRs <- convertmiRs(l$miRs)</pre>
\#colnames(l\$d) \leftarrow c(l\$miRs, l\$mRs)
# 2. Estimate causal effects
nmiR <- length(l$miRs)</pre>
nmR <- length(1$mRs)
standardizedData <- scale(1$d)</pre>
write.table(standardizedData, file = paste(directoryPath, "Lasso_BRCA_input.csv", sep = ""),
            sep = ",", row.names = FALSE)
las=Lasso(paste(directoryPath, "Lasso_BRCA_input.csv", sep = ""), cause = 1:nmiR,
                                    effect = (nmiR+1):(nmiR+nmR))
## Write the result
write.csv(las, file = paste(directoryPath, "Lasso_BRCA_result.csv", sep = ""))
# 3. Validate
# colnames(las) <- convertmiRs(colnames(las))</pre>
# Get the result
las <- read.csv(file = paste(directoryPath, "Lasso_BRCA_result.csv", sep = ""), row.names = 1)</pre>
las <- data.matrix(las)</pre>
row.names(las) <- 1$mRs</pre>
nmiR <- ncol(las)</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
for(topk in c(500, 1000, 1500, 2000)) {
```

```
top = Extopk(las, topk)
  write.csv(top, file = paste(directoryPath, "Lasso_BRCA_top_", topk, ".csv", sep = ""))
  topConfirmed = Validation(top, fileName)
  write.csv(topConfirmed[[1]], file = paste(directoryPath, "Lasso_BRCA_topConfirmed_", topk,
                                            ".csv", sep = ""))
}
##========
# Validate for each miRNA
# colnames(las) <- convertmiRs(colnames(las))</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
for(k in c(50, 100, 150, 200)) {
  topConfirmed = NULL
 top = NULL
 for(i in 1:nmiR) {
   miRiTopk = bRank(las, i, k, FALSE)
   top <- rbind(top, miRiTopk)</pre>
   topConfirmedi = Validation(miRiTopk, fileName)
   if(nrow(topConfirmedi[[1]]) != 0) {
      topConfirmed <- rbind(topConfirmed, topConfirmedi[[1]])</pre>
  write.csv(top, file = paste(directoryPath, "Lasso_BRCA_top_eachmiR_", k, ".csv",
                                       sep = ""), row.names = FALSE)
  write.csv(topConfirmed, file = paste(directoryPath, "Lasso_BRCA_topConfirmed_eachmiR_", k, ".csv",
                                       sep = ""), row.names = FALSE)
}
##========
```

#### Pearson in miRLAB with BRCA

```
# Reset the environment
rm(list = ls ())
library(miRBaseConverter)
library(miRLAB)
##============
# Set environment variables here
directoryPath <- "C:/Users/phavy022/MyDoc/02CausalInferenceMethods/Experiment-Final/InOut/"</pre>
# directoryPath <- "R/CausalCompare/"</pre>
topk_miR <- 30
topk_mR <- 1500
topk <- 15000
k <- 500
source(paste(directoryPath, "CausalCompare.R", sep=""))
# 1. Get data set
load(paste(directoryPath, "01BRCA/BRCA_matchedData2.RData", sep=""))
```

```
# Get significant miRs and mRs from matchedData
1 <- getDatabyMAD(matchedData, topk_miR, topk_mR)</pre>
#l$miRs <- convertmiRs(l$miRs)</pre>
\#colnames(l\$d) \leftarrow c(l\$miRs, l\$mRs)
# 2. Estimate causal effects
nmiR <- length(l$miRs)</pre>
nmR <- length(1$mRs)
standardizedData <- scale(1$d)</pre>
write.table(standardizedData, file = paste(directoryPath, "Pearson_BRCA_input.csv", sep = ""),
            sep = ",", row.names = FALSE)
results=Pearson(paste(directoryPath, "Pearson_BRCA_input.csv", sep = ""), cause = 1:nmiR,
                                     effect = (nmiR+1):(nmiR+nmR))
## Write the result
write.csv(results, file = paste(directoryPath, "Pearson_BRCA_result.csv", sep = ""))
# 3. Validate
# colnames(results) <- convertmiRs(colnames(results))</pre>
# Get the result
results <- read.csv(file = paste(directoryPath, "Pearson_BRCA_result.csv", sep = ""), row.names = 1)
results <- data.matrix(results)</pre>
row.names(results) <- 1$mRs</pre>
nmiR <- ncol(results)</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
for(topk in c(500, 1000, 1500, 2000)) {
  top = Extopk(results, topk)
  write.csv(top, file = paste(directoryPath, "Pearson_BRCA_top_", topk, ".csv", sep = ""))
  topConfirmed = Validation(top, fileName)
  write.csv(topConfirmed[[1]], file = paste(directoryPath, "Pearson_BRCA_topConfirmed_",
                                              topk, ".csv", sep = ""))
}
##=========
# Validate for each miRNA
# colnames(results) <- convertmiRs(colnames(results))</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
for(k in c(50, 100, 150, 200)) {
  topConfirmed = NULL
  top = NULL
  for(i in 1:nmiR) {
    miRiTopk = bRank(results, i, k, FALSE)
    top <- rbind(top, miRiTopk)</pre>
    topConfirmedi = Validation(miRiTopk, fileName)
    if(nrow(topConfirmedi[[1]]) != 0) {
      topConfirmed <- rbind(topConfirmed, topConfirmedi[[1]])</pre>
    }
  }
```

## Borda for hiddenICP 1/3, Pearson and Lasso with BRCA

```
# Reset the environment
rm(list = ls ())
library(miRBaseConverter)
library(miRLAB)
library(robustbase)
library(pcalg)
library(Rgraphviz)
library(CancerSubtypes)
# Set environment variables here
directoryPath <- "C:/Users/phavy022/MyDoc/02CausalInferenceMethods/Experiment-Final/InOut/"
# directoryPath <- "R/CausalCompare/"</pre>
source(paste(directoryPath, "CausalCompare.R", sep=""))
# 1. Get the results hiddenICP, Pearson and Lasso
hid <- read.csv(paste(directoryPath, "hiddenICP_BRCA13_result.csv", sep = ""), row.names = 1)
pea <- read.csv(paste(directoryPath, "Pearson_BRCA_result.csv", sep = ""), row.names = 1)</pre>
las <- read.csv(paste(directoryPath, "Lasso_BRCA_result.csv", sep = ""), row.names = 1)</pre>
hid <- data.matrix(hid)
pea <- data.matrix(pea)</pre>
las <- data.matrix(las)</pre>
row.names(pea) <- row.names(hid)</pre>
row.names(las) <- row.names(hid)
# 2. Borda
borda=Borda(list(hid, pea, las))
## Write the result
write.csv(borda, file = paste(directoryPath, "Borda_hiddenICP_13_BRCA_result.csv", sep = ""))
# colnames(borda) <- convertmiRs(colnames(borda))</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
for(topk in c(500, 1000, 1500, 2000)) {
 top = Extopk(borda, topk)
 write.csv(top, file = paste(directoryPath, "Borda_hiddenICP_13_BRCA_top_", topk, ".csv", sep = ""))
```

```
topConfirmed = Validation(top, fileName)
  write.csv(topConfirmed[[1]], file = paste(directoryPath, "Borda_hiddenICP_13_BRCA_topConfirmed_",
                                             topk, ".csv", sep = ""))
}
##========
# Validate for each miRNA
# colnames(borda) <- convertmiRs(colnames(borda))</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
nmiR <- ncol(hid)</pre>
for(k in c(50, 100, 150, 200)) {
  topConfirmed = NULL
  top = NULL
  for(i in 1:nmiR) {
    miRiTopk = bRank(borda, i, k, FALSE)
    top <- rbind(top, miRiTopk)</pre>
    topConfirmedi = Validation(miRiTopk, fileName)
    if(nrow(topConfirmedi[[1]]) != 0) {
      topConfirmed <- rbind(topConfirmed, topConfirmedi[[1]])</pre>
    }
  }
  write.csv(top, file = paste(directoryPath, "Borda_hiddenICP_13_BRCA_top_eachmiR_", k, ".csv",
                                       sep = ""), row.names = FALSE)
  write.csv(topConfirmed, file = paste(directoryPath, "Borda_hiddenICP_13_BRCA_topConfirmed_eachmiR_",
                                       k, ".csv",
                                       sep = ""), row.names = FALSE)
}
##========
```

#### Borda for hiddenICP PAM50, Pearson and Lasso with BRCA

```
# Reset the environment
rm(list = ls ())
library(miRBaseConverter)
library(miRLAB)
library(robustbase)
library(pcalg)
library(Rgraphviz)
library(CancerSubtypes)
# Set environment variables here
directoryPath <- "C:/Users/phavy022/MyDoc/02CausalInferenceMethods/Experiment-Final/InOut/"
# directoryPath <- "R/CausalCompare/"</pre>
source(paste(directoryPath, "CausalCompare.R", sep=""))
# 1. Get the results hiddenICP, Pearson and Lasso
hid <- read.csv(paste(directoryPath, "hiddenICP_BRCA_pam50_result.csv", sep = ""), row.names = 1)
```

```
pea <- read.csv(paste(directoryPath, "Pearson_BRCA_result.csv", sep = ""), row.names = 1)</pre>
las <- read.csv(paste(directoryPath, "Lasso_BRCA_result.csv", sep = ""), row.names = 1)</pre>
hid <- data.matrix(hid)</pre>
pea <- data.matrix(pea)</pre>
las <- data.matrix(las)</pre>
row.names(pea) <- row.names(hid)
row.names(las) <- row.names(hid)
# 2. Borda
borda=Borda(list(hid, pea, las))
## Write the result
write.csv(borda, file = paste(directoryPath, "Borda_hiddenICP_pam50_BRCA_result.csv", sep = ""))
# 3. Validate
# colnames(borda) <- convertmiRs(colnames(borda))</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")
for(topk in c(500, 1000, 1500, 2000)) {
  top = Extopk(borda, topk)
  write.csv(top, file = paste(directoryPath, "Borda_hiddenICP_pam50_BRCA_top_", topk, ".csv",
                               sep = ""))
 topConfirmed = Validation(top, fileName)
  write.csv(topConfirmed[[1]], file = paste(directoryPath, "Borda_hiddenICP_pam50_BRCA_topConfirmed_",
                                              topk, ".csv", sep = ""))
}
##========
# Validate for each miRNA
# colnames(borda) <- convertmiRs(colnames(borda))</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
nmiR <- ncol(hid)
for(k in c(50, 100, 150, 200)) {
  topConfirmed = NULL
  top = NULL
 for(i in 1:nmiR) {
    miRiTopk = bRank(borda, i, k, FALSE)
    top <- rbind(top, miRiTopk)</pre>
    topConfirmedi = Validation(miRiTopk, fileName)
    if(nrow(topConfirmedi[[1]]) != 0) {
      topConfirmed <- rbind(topConfirmed, topConfirmedi[[1]])</pre>
    }
  write.csv(top, file = paste(directoryPath, "Borda_hiddenICP_pam50_BRCA_top_eachmiR_", k, ".csv",
                                        sep = ""), row.names = FALSE)
  write.csv(topConfirmed, file = paste(directoryPath,
                                        "Borda_hiddenICP_pam50_BRCA_topConfirmed_eachmiR_",
                                        k, ".csv",
                                        sep = ""), row.names = FALSE)
}
```

## Borda for idaFast, Pearson and Lasso with BRCA

```
# Reset the environment
rm(list = ls ())
library(miRBaseConverter)
library(miRLAB)
library(robustbase)
library(pcalg)
library(Rgraphviz)
library(CancerSubtypes)
# Set environment variables here
directoryPath <- "C:/Users/phavy022/MyDoc/02CausalInferenceMethods/Experiment-Final/InOut/"
# directoryPath <- "R/CausalCompare/"</pre>
source(paste(directoryPath, "CausalCompare.R", sep=""))
# 1. Get the results idaFast, Pearson and Lasso
ida <- read.csv(paste(directoryPath, "pc_idaFast_BRCA_result.csv", sep = ""), row.names = 1)
pea <- read.csv(paste(directoryPath, "Pearson_BRCA_result.csv", sep = ""), row.names = 1)</pre>
las <- read.csv(paste(directoryPath, "Lasso_BRCA_result.csv", sep = ""), row.names = 1)</pre>
ida <- data.matrix(ida)</pre>
pea <- data.matrix(pea)</pre>
las <- data.matrix(las)</pre>
row.names(pea) <- row.names(ida)</pre>
row.names(las) <- row.names(ida)
# 2. Borda
borda=Borda(list(ida, pea, las))
## Write the result
write.csv(borda, file = paste(directoryPath, "Borda_idafast_BRCA_result.csv", sep = ""))
# 3. Validate
# colnames(borda) <- convertmiRs(colnames(borda))</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
for(topk in c(500, 1000, 1500, 2000)) {
  top = Extopk(borda, topk)
  write.csv(top, file = paste(directoryPath, "Borda_idaFast_BRCA_top_", topk, ".csv", sep = ""))
 topConfirmed = Validation(top, fileName)
  write.csv(topConfirmed[[1]], file = paste(directoryPath, "Borda_idaFast_BRCA_topConfirmed_",
                                            topk, ".csv", sep = ""))
}
##=========
# Validate for each miRNA
# colnames(borda) <- convertmiRs(colnames(borda))</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
```

```
nmiR <- ncol(ida)</pre>
for(k in c(50, 100, 150, 200)) {
  topConfirmed = NULL
  top = NULL
  for(i in 1:nmiR) {
   miRiTopk = bRank(borda, i, k, FALSE)
   top <- rbind(top, miRiTopk)</pre>
   topConfirmedi = Validation(miRiTopk, fileName)
    if(nrow(topConfirmedi[[1]]) != 0) {
      topConfirmed <- rbind(topConfirmed, topConfirmedi[[1]])</pre>
   }
  }
  write.csv(top, file = paste(directoryPath, "Borda_idaFast_BRCA_top_eachmiR_", k, ".csv",
                                        sep = ""), row.names = FALSE)
  write.csv(topConfirmed, file = paste(directoryPath, "Borda_idaFast_BRCA_topConfirmed_eachmiR_",
                                       k, ".csv",
                                        sep = ""), row.names = FALSE)
}
##=========
```

# Borda for jointIDA, Pearson and Lasso with BRCA

```
# Reset the environment
rm(list = ls ())
library(miRBaseConverter)
library(miRLAB)
library(robustbase)
library(pcalg)
library(Rgraphviz)
library(CancerSubtypes)
# Set environment variables here
directoryPath <- "C:/Users/phavy022/MyDoc/02CausalInferenceMethods/Experiment-Final/InOut/"
# directoryPath <- "R/CausalCompare/"</pre>
source(paste(directoryPath, "CausalCompare.R", sep=""))
# 1. Get the results jointIDA, Pearson and Lasso
joi <- read.csv(paste(directoryPath, "jointIDA_direct_BRCA_result.csv", sep = ""), row.names = 1)
pea <- read.csv(paste(directoryPath, "Pearson_BRCA_result.csv", sep = ""), row.names = 1)</pre>
las <- read.csv(paste(directoryPath, "Lasso_BRCA_result.csv", sep = ""), row.names = 1)</pre>
joi <- data.matrix(joi)</pre>
pea <- data.matrix(pea)</pre>
las <- data.matrix(las)</pre>
row.names(pea) <- row.names(joi)</pre>
row.names(las) <- row.names(joi)</pre>
```

```
# 2. Borda
borda=Borda(list(joi, pea, las))
## Write the result
write.csv(borda, file = paste(directoryPath, "Borda_jointIDA_BRCA_result.csv", sep = ""))
# 3. Validate
# colnames(borda) <- convertmiRs(colnames(borda))</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
for(topk in c(500, 1000, 1500, 2000)) {
  top = Extopk(borda, topk)
  write.csv(top, file = paste(directoryPath, "Borda_jointIDA_BRCA_top_", topk, ".csv", sep = ""))
  topConfirmed = Validation(top, fileName)
  write.csv(topConfirmed[[1]], file = paste(directoryPath, "Borda_jointIDA_BRCA_topConfirmed_",
                                             topk, ".csv", sep = ""))
}
##========
# Validate for each miRNA
# colnames(borda) <- convertmiRs(colnames(borda))</pre>
fileName <- paste(directoryPath, "groundtruth.csv", sep = "")</pre>
nmiR <- ncol(joi)</pre>
for(k in c(50, 100, 150, 200)) {
  topConfirmed = NULL
  top = NULL
  for(i in 1:nmiR) {
    miRiTopk = bRank(borda, i, k, FALSE)
    top <- rbind(top, miRiTopk)</pre>
    topConfirmedi = Validation(miRiTopk, fileName)
    if(nrow(topConfirmedi[[1]]) != 0) {
      topConfirmed <- rbind(topConfirmed, topConfirmedi[[1]])</pre>
    }
  }
  write.csv(top, file = paste(directoryPath, "Borda_jointIDA_BRCA_top_eachmiR_", k, ".csv",
                                        sep = ""), row.names = FALSE)
  write.csv(topConfirmed, file = paste(directoryPath, "Borda_jointIDA_BRCA_topConfirmed_eachmiR_",
                                        k, ".csv",
                                        sep = ""), row.names = FALSE)
}
##=========
```