PSB 2017: Exploring the Reproducibility of Probabilistic Causal Molecular Network Models

R Code used in analyses and generating figures and tables

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All files required to reproduce the results presented in this paper are located in the following directory:

gitPath <- c("https://raw.githubusercontent.com/divara01/PSB2017_ReproducibilityOfBNs/")</pre>

Create Edges_all object:

This code breaks up the results from the RIMBAnet algorithm to save the resulting edges into a dataframe. Singletons are removed. Each edge is associated with a resulting posterior probability which is output from the algorithm as well. The resulting edges_all file is saved in the current working directory. This is where all downstream output will be saved.

```
options(stringsAsFactors = F)
rm(list = ls())
#### Load all files into R####
gitPath <- c("https://raw.githubusercontent.com/divara01/PSB2017_ReproducibilityOfBNs/")</pre>
dataset <- c("GTEx", "STARNET", "Simulation")</pre>
subSets \leftarrow c(100, 90, 80, 50, 10)
replicates <- c(1:5)
FullLabels <- as.vector(sapply(subSets, function(x) sapply(replicates,</pre>
    function(y) paste(x, y, sep = "_"))))
for (datasetIndex in 1:length(dataset)) {
    if (datasetIndex < 3) {</pre>
        labels <- FullLabels[-grep("10_", FullLabels)]</pre>
    } else {
        labels <- FullLabels
    }
    for (index in 1:length(labels)) {
        filename <- paste(gitPath, "data/", dataset[datasetIndex],</pre>
            "_", labels[index], "_EdgeFile", sep = "")
        edgeFile_i <- read.csv(filename, quote = "\"", sep = "\t",</pre>
            comment.char = "", stringsAsFactors = FALSE)
        edgeFile i <- as.data.frame.matrix(t(apply(edgeFile i,</pre>
            1, function(x) {
                 strsplit(x, "->", fixed = T)[[1]]
            })))
        colnames(edgeFile_i) <- c("FromNode", "ToNode")</pre>
        PostProbFile_i <- read.csv(paste(gitPath, "data/", dataset[datasetIndex],</pre>
            "_", labels[index], "_labeledEdges", sep = ""), quote = "\"",
            comment.char = "", stringsAsFactors = FALSE, header = F)
        PostProbFile_i$Edge <- unlist(lapply(strsplit(PostProbFile_i$V1,</pre>
            " "), function(x) x[1]))
        PostProbFile_i$PostProb <- as.numeric(unlist(lapply(strsplit(PostProbFile_i$V1,
            " [[]label="), function(x) lapply(strsplit(x[2],
            "[]];"), function(y) y[1]))))
        assign(paste(dataset[datasetIndex], labels[index], "EdgeFile",
            sep = "_"), edgeFile_i)
        assign(paste(dataset[datasetIndex], "PostProbFile", labels[index],
            sep = "_"), PostProbFile_i)
    }
rm(index, edgeFile_i, PostProbFile_i)
for (datasetIndex in 1:length(dataset)) {
    message(paste(dataset[datasetIndex], "running...", sep = " "))
    edges_all <- data.frame(matrix(NA, nrow = 1, ncol = 5))</pre>
    colnames(edges_all) <- c("node1", "node2", "postProb", "repRun",</pre>
        "data_type")
```

```
if (datasetIndex < 3) {</pre>
    labels <- FullLabels[-grep("10_", FullLabels)]</pre>
    labels <- FullLabels
}
for (index in 1:length(labels)) {
    edgeFile i <- get(paste(dataset[datasetIndex], labels[index],</pre>
        "EdgeFile", sep = " "))
    edge_i <- paste(edgeFile_i$FromNode, edgeFile_i$ToNode,</pre>
        sep = "->")
    PostProbFile_i <- get(paste(dataset[datasetIndex], "PostProbFile",</pre>
        labels[index], sep = "_"))
    postProb <- as.character(substr(PostProbFile_i$PostProb[match(edge_i,</pre>
        PostProbFile_i$Edge)], 1, 3))
    node1 <- as.vector(unlist(lapply(strsplit(edge_i, "->"),
        function(x) x[1])))
    node2 <- as.vector(unlist(lapply(strsplit(edge_i, "->"),
        function(x) x[2])))
    repRun <- rep(as.vector(strsplit(labels[index], "_")[[1]][2]),</pre>
        length(edge_i))
    data_type <- rep(as.vector(strsplit(labels[index], "_")[[1]][1]),</pre>
        length(edge_i))
    vect <- cbind(node1, node2, postProb, repRun, data_type)</pre>
    edges_all <- rbind.data.frame(edges_all, vect)</pre>
edges_all <- edges_all[-1, ]
# Add the additional rows for all posterior probabilities
# below the actual value (so for postProb = 0.5, add row for
# 0.4, 0.3, 0.2, and 0.1)
postProbOptions \leftarrow seq(0.1, 1, 0.1)
for (index in 1:nrow(edges_all)) {
    postProb_i <- edges_all$postProb[index]</pre>
    pos <- which(postProbOptions == postProb_i)</pre>
    tempEdgeFile <- c()</pre>
    if (pos != 1) {
        for (i in 1:(pos - 1)) {
             tempEdgeFile <- rbind(tempEdgeFile, edges_all[index,</pre>
             tempEdgeFile$postProb[nrow(tempEdgeFile)] <- postProbOptions[i]</pre>
        edges_all <- rbind.data.frame(edges_all, tempEdgeFile)</pre>
    }
}
edges_all <- edges_all[order(as.numeric(edges_all$data_type),</pre>
    decreasing = TRUE), ]
edgesAll_filename <- paste(dataset[datasetIndex], "_edges_all_RIMBAnet.RData",</pre>
    sep = "")
save(edges_all, file = edgesAll_filename)
message(paste(dataset[datasetIndex], "complete!", sep = " "))
```

}

Analysis Set-up:

```
options(stringsAsFactors = F)
rm(list = ls())
library(ggplot2)
library(reshape)
library(RCurl)
```

Load Functions

summarySE Function

This function provides the count, mean, standard deviation, standard error of the mean, and confidence interval (default 95%). The inputs for the functions are the following:

- data: a data frame.
- measurevar: the name of a column that contains the variable to be summariezed
- groupvars: a vector containing names of columns that contain grouping variables
- na.rm: a boolean that indicates whether to ignore NA's
- conf.interval: the percent range of the confidence interval (default is 95%)

```
summarySE <- function(data = NULL, measurevar, groupvars = NULL,</pre>
   na.rm = FALSE, conf.interval = 0.95, .drop = TRUE) {
   library(plyr)
    # New version of length which can handle NA's: if na.rm==T,
    # don't count them
   length2 <- function(x, na.rm = FALSE) {</pre>
        if (na.rm)
            sum(!is.na(x)) else length(x)
   }
    # This does the summary. For each group's data frame, return
    \# a vector with N, mean, and sd
   datac <- ddply(data, groupvars, .drop = .drop, .fun = function(xx,</pre>
        col) {
        c(N = length2(xx[[col]], na.rm = na.rm), mean = mean(xx[[col]],
            na.rm = na.rm), sd = sd(xx[[col]], na.rm = na.rm))
   }, measurevar)
    # Rename the 'mean' column
   datac <- rename(datac, c(mean = measurevar))</pre>
   datac$se <- datac$sd/sqrt(datac$N) # Calculate standard error of the mean
    # Confidence interval multiplier for standard error Calculate
    # t-statistic for confidence interval: e.g., if conf.interval
    # is .95, use .975 (above/below), and use df=N-1
```

```
ciMult <- qt(conf.interval/2 + 0.5, datac$N - 1)
datac$ci <- datac$se * ciMult

return(datac)
}</pre>
```

get_ABoverA_directed (Figure 2A)

Given two networks A and B, obtain the fraction of edges in both A and B over the number of edges in A assuming fixed probability for edges in A and any edge in B.

```
get_ABoverA_directed <- function(edges_all, A_name, B_name, prob,</pre>
    sameProb = F) {
    if (A_name == B_name) {
        if (sameProb) {
            aANDb = length(intersect(edges_all$edge_name[edges_all$postProb ==
                prob & edges_all$type_name == A_name], edges_all$edge_name[edges_all$postProb ==
                prob & edges_all$type_name != B_name]))
        } else {
            aANDb = length(intersect(edges_all$edge_name[edges_all$postProb ==
                prob & edges_all$type_name == A_name], edges_all$edge_name[edges_all$type_name !=
                B name]))
        }
   } else {
        if (sameProb) {
            aANDb = length(intersect(edges_all$edge_name[edges_all$postProb ==
                prob & edges_all$type_name == A_name], edges_all$edge_name[edges_all$postProb ==
                prob & edges_all$type_name == B_name]))
        } else {
            aANDb = length(intersect(edges_all$edge_name[edges_all$postProb ==
                prob & edges_all$type_name == A_name], edges_all$edge_name[edges_all$type_name ==
                B_name]))
        }
    a = sum(edges_all$postProb == prob & edges_all$type_name ==
        A name)
   return(aANDb/a)
}
```

Load in Data

```
# Load in the data:
load("STARNET_edges_all_RIMBAnet.RData")
edges_all_starnet = edges_all
rm(edges_all)
load("GTEx_edges_all_RIMBAnet.RData")
edges_all_gtex = edges_all
rm(edges_all)

## adding edge name:
edges_all_starnet$edge_name = paste(edges_all_starnet$node1,
```

```
edges_all_starnet$node2, sep = "_")
edges_all_gtex$edge_name = paste(edges_all_gtex$node1, edges_all_gtex$node2,
    sep = "_")
## adding type_name:
edges_all_starnet$type_name = paste(edges_all_starnet$data_type,
    edges_all_starnet$repRun, sep = "_")
edges_all_gtex$type_name = paste(edges_all_gtex$data_type, edges_all_gtex$repRun,
    sep = "_")
```

Analyses for Figures

Figure 2A: Precision/Recall Plots for edges

Network A: Subsampling Network

Network B: Complete Network (100% Replicate 1)

Precision = $A \cap B / A$

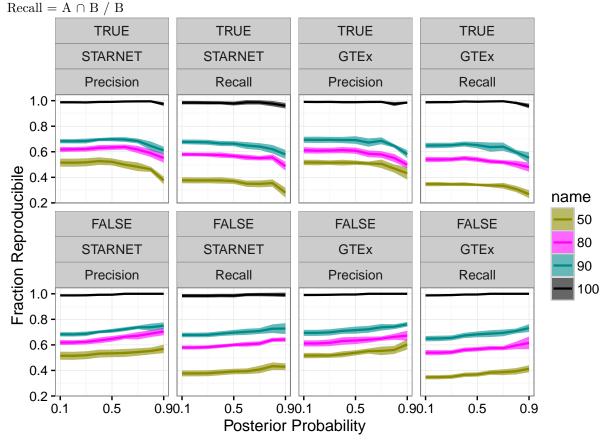


Figure 2B: Evaluation of Path Length

This code will prepare the data. It is highly advised that the code is run and resulting object is saved as it takes some time to run.

```
gtex_fn = "GTEx_edges_all_RIMBAnet.RData"
```

```
load(gtex_fn)
head(edges_all)
real_network_all = edges_all[edges_all$repRun == 1 & edges_all$data_type ==
    100, ]
dim(real_network_all)
threads = 18
library(foreach)
library(parallel)
library(doMC)
options(cores = detectCores())
registerDoMC(threads)
path <- getwd()</pre>
all_out = c()
tests = cbind(rep(1:9, 15), c(rep(1, 5 * 9), rep(2, 5 * 9), rep(3, 5))
    5 * 9)), rep(1:5))
colnames(tests) = c("prob", "subsampling_idx", "repRun")
edgesInPath <- function(edgeList1, g2, path_length, directed = TRUE) {</pre>
    if (!directed) {
        g2 = as.undirected(g2)
    }
    InPath = apply(edgeList1, 1, function(x) {
        n1 = x[1]
        n2 = x[2]
        if (sum(is.element(c(n1, n2), V(g2)\$name)) == 2) {
            p = get.shortest.paths(g2, n1, n2)
            1 = min(unlist(lapply(p$vpath, length)))
            if (1 <= (path_length + 1) & 1 != 0) {
                out = 1
            } else {
                out = 0
        } else {
            out = 0
        }
        out
    })
    return(InPath)
}
compare_twoBN <- function(rand1, rand2, maxK = 10, text = TRUE) {</pre>
    g1 <- graph.edgelist(as.matrix(rand1))</pre>
    g2 <- graph.edgelist(as.matrix(rand2))</pre>
```

```
rownames(rand1) = paste(rand1[, 1], rand1[, 2], sep = "_")
    rownames(rand2) = paste(rand2[, 1], rand2[, 2], sep = "_")
    out = c()
    for (i in 1:maxK) {
        if (text) {
            cat("on run path of", i, "\n")
        e1_in_g2 = edgesInPath(as.matrix(rand1), g2, i)
        e2_in_g1 = edgesInPath(as.matrix(rand2), g1, i)
        fracE1_in_G2 = sum(e1_in_g2/length(e1_in_g2))
        fracE2_in_G1 = sum(e2_in_g1/length(e2_in_g1))
        e1_in_g2_undir = edgesInPath(as.matrix(rand1), g2, i,
            FALSE)
        e2_in_g1_undir = edgesInPath(as.matrix(rand2), g1, i,
            FALSE)
        fracE1_in_G2_undir = sum(e1_in_g2_undir/length(e1_in_g2_undir))
        fracE2_in_G1_undir = sum(e2_in_g1_undir/length(e2_in_g1_undir))
        out = rbind(out, cbind(fracE1_in_G2, fracE2_in_G1, fracE1_in_G2_undir,
            fracE2_in_G1_undir))
    }
    rownames(out) = paste("path_", 1:maxK, sep = "")
    return(out)
}
run_Network_comparisions <- function(edges_all, real_network,</pre>
    probs = seq(0.1, 0.9, 0.1)) {
    all = c()
    for (prob in probs) {
        net = edges_all[edges_all$postProb == prob, c("node1",
            "node2")]
        comp = compare_twoBN(net, real_network, maxK = 10, text = F)
        comp = as.data.frame.matrix(comp)
        comp$post_prob_net1 = prob
        comp$path = 1:nrow(comp)
        all = rbind(all, comp)
    cat("\n")
    return(all)
}
paths = c(50, 80, 90)
## In parallel calculating all the different comparisions:
all_out <- foreach(i = 1:nrow(tests), combine = rbind) %dopar%</pre>
    {
        print(paste("real network", i, "start time:", Sys.time()))
```

We are not displaying the figure here, however, below is the code to generate Figure 2B:

```
load("Fig2B GTEx updated.RData")
results$name = sapply(results$Name, function(x) {
   paste(strsplit(x, "_")[[1]][1], collapse = "_")
})
results$repRun = sapply(results$Name, function(x) {
    paste(strsplit(x, "_")[[1]][2], collapse = "_")
})
results = results[, colnames(results) != "Name"]
results = results[, colnames(results) != "fracE1_in_G2_undir"]
results = results[, colnames(results) != "fracE2_in_G1_undir"]
melted_data = melt(results, id = c("name", "path", "post_prob_net2",
    "post_prob_net1", "repRun"))
melted_data$path = as.factor(melted_data$path)
melted_data$post_prob_net1_name = paste("SubSampling Posterior: ",
    melted data$post prob net1, sep = "")
melted_data$post_prob_net2_name = paste("Real Posterior: ", melted_data$post_prob_net2,
    sep = "")
ggplotDF = melted_data[melted_data$variable == "fracE1_in_G2" &
    is.element(melted_data$post_prob_net1_name, c("SubSampling Posterior: 0.1",
        "SubSampling Posterior: 0.5", "SubSampling Posterior: 0.9")),
   ]
ggplotDF2 = ggplotDF[is.element(ggplotDF$post_prob_net2, c("0.1",
    "0.5", "0.9")), ]
ggplotDF2_summary <- summarySE(ggplotDF2, "value", groupvars = c("name",</pre>
    "path", "post_prob_net1", "post_prob_net2"))
ggplotDF2_summary$path = as.numeric(as.character(ggplotDF2_summary$path))
ggplotDF2_summary$post_prob_net1_name = as.character(ggplotDF2_summary$post_prob_net1)
ggplotDF2_summary$post_prob_net2_name = as.character(ggplotDF2_summary$post_prob_net2)
fig_2b_gtex <- ggplot(ggplotDF2_summary, aes(path)) + geom_line(aes(y = value,
```

```
colour = post_prob_net2_name)) + geom_ribbon(aes(ymin = value -
sd, ymax = value + sd, fill = post_prob_net2_name), alpha = 0.8) +
facet_wrap(~name + post_prob_net1_name, ncol = 3) + xlab("Path Length") +
scale_color_manual(values = c("orange", "green", "grey50")) +
scale_fill_manual(values = c("orange", "green", "grey50")) +
ggtitle("Fraction of SubSampling Edges in Real BN at Different SubSamping Thresholds") +
theme(axis.text.x = element_text(size = 12), title = element_text(size = 16)) +
theme_bw() + scale_x_continuous(breaks = 1:10)
show(fig_2b_gtex)
```

Figure 2C: Clique Associates for Various Correlation Cutoffs for Edges

Create Edge files based on correlation thresholds

```
dataFlag <- c("GTEx", "STARNET")</pre>
subSets \leftarrow c(100, 90, 80, 50)
replicates <- c(1:5)
labels <- as.vector(sapply(subSets, function(x) sapply(replicates,</pre>
    function(y) paste(x, y, sep = "_"))))
dataFN <- as.vector(sapply(dataFlag, function(x) sapply(labels,</pre>
    function(y) paste(x, y, sep = "_"))))
for (labelsTagIndex in 1:length(dataFN)) {
    labelsTag <- dataFN[labelsTagIndex]</pre>
    for (cutoff in c(0.1, 0.05, 0.01)) {
        message(paste(dataSet, "-", cutoff, "-", labelsTag, "running..."))
        data = read.table(paste(gitPath, "RIMBAnetInput/", dataSet,
            "_", labelsTag, "_BC.txt", sep = ""), quote = "\"",
            comment.char = "", stringsAsFactors = FALSE, row.names = 1)
        cor_data = cor(t(data))^2
        rsqrd_cutoff = quantile(unlist(cor_data[upper.tri(cor_data)]),
            1 - cutoff)
        keep_edges = which(cor_data >= rsqrd_cutoff, arr.ind = T)
        edges = t(apply(keep_edges, 1, function(X) {
            t = c(rownames(cor_data)[X[1]], rownames(cor_data)[X[2]])
            t[order(t)]
        }))
        edges = unique(edges)
        edges = edges[edges[, 1] != edges[, 2], ]
        write.table(edges, sep = "\t", quote = F, col.names = F,
            row.names = F, file = paste(dataSet, labelsTag, "_",
                cutoff, "_topCorEdges.txt", sep = ""))
    }
```

We feed the resulting edge files into the COS program to determine which edges have both nodes in a single clique. The code below parses the results from COS and saves the cliques into a list for each dataset, GTEx and STARNET.

```
subSets \leftarrow c(100, 90, 80, 50)
replicates <- c(1:5)
postProbs <- c(seq(0.1, 0.9, 0.1))
thresholds \leftarrow c(0.01, 0.05, 0.1)
datasets <- c("GTEx", "STARNET")</pre>
labels <- as.vector(sapply(subSets, function(x) sapply(replicates,</pre>
    function(y) sapply(thresholds, function(z) paste(x, y, z,
        sep = "_")))))
for (datasetIndex in 1:length(datasets)) {
    message(paste(datasets[datasetIndex], "running..."))
    MappedCliques_i <- vector("list", length(labels))</pre>
    for (index in 1:length(labels)) {
        cliqueMap_i <- read.table(paste(datasets[datasetIndex],</pre>
            "_", labels[index], "_topCorEdges.txt.map", sep = ""),
            quote = "\"", comment.char = "")
        filename <- paste(datasets[datasetIndex], labels[index],</pre>
            "MappedCliques", sep = " ")
        if (file.exists(paste(datasets[datasetIndex], "_", labels[index],
            "_topCorEdges.txt.mcliques_3_communities.txt", sep = ""))) {
            cliqueCommunity_i <- read.table(paste(datasets[datasetIndex],</pre>
                "_", labels[index], "_topCorEdges.txt.mcliques_3_communities.txt",
                sep = ""), quote = "\"", comment.char = "", fill = TRUE)
            # First column is not delimited properly, so need to split
            # the label out:
            cliqueCommunity i$V1 <- as.integer(as.vector(unlist(lapply(strsplit(cliqueCommunity i$V1,
                ":"), function(x) x[2]))))
            mappedCliques <- apply(cliqueCommunity_i, 2, function(x) cliqueMap_i$V1[match(x,</pre>
                cliqueMap i$V2)])
            # Store the mapped cliques locally in R workspace
            assign(filename, mappedCliques)
            # Store the mapped cliques into a list
            MappedCliques_i[[index]] <- mappedCliques</pre>
        names(MappedCliques_i)[[index]] <- filename</pre>
    }
    # Save RData file
    save(MappedCliques_i, file = paste(datasets[datasetIndex],
        "_MappedCliquesList.RData", sep = ""))
    message(paste(datasets[datasetIndex], "Complete!"))
```

Once cliques are called, add a flag for each edge as to whether both nodes are found in the same clique. The results from this code should be saved and then loaded back in.

```
options(stringsAsFactors = F)
rm(list = ls())
library(ggplot2)
library(reshape)
gitPath <- c("https://raw.githubusercontent.com/divara01/PSB2017_ReproducibilityOfBNs/")</pre>
inclique <- function(Clique_list, name, node1, node2) {</pre>
    cliques <- Clique_list[[name]]</pre>
    if (class(cliques) == "character") {
        cliques = matrix(cliques, nrow = 1)
    }
    if (length(cliques) == 0) {
        cat("this run has NO cliques,", name, "\n")
        return(FALSE)
    pos n1 <- which(!is.na(apply(cliques, 1, function(x) match(node1,</pre>
        x))))
    pos_n2 <- which(!is.na(apply(cliques, 1, function(x) match(node2,</pre>
        x))))
    cliquesContainingBothNodes <- intersect(pos n1, pos n2)</pre>
    if (length(cliquesContainingBothNodes) > 0) {
        return(TRUE)
    } else {
        return(FALSE)
    }
}
## loading Clique List
load(paste(gitPath, "COSresults/STARNET_MappedCliquesList.RData",
    sep = ""))
## loading edges_all
load("STARNET edges all RIMBAnet.RData")
CompleteCliqueFlag <- data.frame(matrix(NA, nrow = nrow(edges_all),</pre>
    ncol = 3)
colnames(CompleteCliqueFlag) <- c("0.01", "0.05", "0.1")</pre>
for (cutOffIndex in 1:ncol(CompleteCliqueFlag)) {
    print(paste(colnames(CompleteCliqueFlag)[cutOffIndex], " Running...",
        sep = "")
    CompleteCliqueFlag[, cutOffIndex] <- sapply(1:nrow(edges_all),</pre>
        function(x) {
            MappedName = paste("STARNET", edges_all$data_type[x],
                edges_all$repRun[x], colnames(CompleteCliqueFlag)[cutOffIndex],
                 "MappedCliques", sep = "_")
            inclique(MappedCliques_i, MappedName, edges_all$node1[x],
                edges_all$node2[x])
        })
```

```
print(paste(colnames(CompleteCliqueFlag)[cutOffIndex], " Complete!",
        sep = ""))
}
colnames(CompleteCliqueFlag) <- paste("inClique", colnames(CompleteCliqueFlag),</pre>
edges_all <- cbind.data.frame(edges_all, CompleteCliqueFlag)</pre>
edges all\sinClique 0.1 <- as.integer(edges all\sinClique 0.1 +
edges_all$inClique_0.05 <- as.integer(edges_all$inClique_0.05 +</pre>
edges all$inClique 0.01 <- as.integer(edges all$inClique 0.01 +
    0)
save(edges_all, file = "STARNET_edges_all_RIMBAnet_cliqueAdded.RData")
## GTEx:
load(paste(gitPath, "COSresults/GTEx_MappedCliquesList.RData",
    sep = ""))
load("GTEx_edges_all_RIMBAnet.RData")
CompleteCliqueFlag <- data.frame(matrix(NA, nrow = nrow(edges_all),</pre>
    ncol = 3))
colnames(CompleteCliqueFlag) <- c("0.01", "0.05", "0.1")</pre>
for (cutOffIndex in 1:ncol(CompleteCliqueFlag)) {
    print(paste(colnames(CompleteCliqueFlag)[cutOffIndex], " Running...",
        sep = "")
    CompleteCliqueFlag[, cutOffIndex] <- sapply(1:nrow(edges_all),</pre>
        function(x) {
            MappedName = paste("GTEx", edges_all$data_type[x],
                edges_all$repRun[x], colnames(CompleteCliqueFlag)[cutOffIndex],
                 "MappedCliques", sep = "_")
            inclique(MappedCliques_i, MappedName, edges_all$node1[x],
                edges_all$node2[x])
        })
    print(paste(colnames(CompleteCliqueFlag)[cutOffIndex], " Complete!",
        sep = "")
}
colnames(CompleteCliqueFlag) <- paste("inClique", colnames(CompleteCliqueFlag),</pre>
    sep = " ")
edges all <- cbind.data.frame(edges all, CompleteCliqueFlag)</pre>
edges_all$inClique_0.1 <- as.integer(edges_all$inClique_0.1 +</pre>
edges_all$inClique_0.05 <- as.integer(edges_all$inClique_0.05 +</pre>
edges_all$inClique_0.01 <- as.integer(edges_all$inClique_0.01 +</pre>
    0)
save(edges_all, file = "GTEx_edges_all_RIMBAnet_cliqueAdded.RData")
```

We are not displaying the figure here, however, below is the code to generate Figure 2C:

```
load("STARNET edges all RIMBAnet cliqueAdded.RData")
edges all starnet = edges all
rm(edges all)
load("GTEx_edges_all_RIMBAnet_cliqueAdded.RData")
edges all gtex = edges all
edges_all_gtex$inClique_0.1 = edges_all_gtex$inClique_0.1 + 0
rm(edges all)
edges_all_starnet$edge_name = paste(edges_all_starnet$node1,
    edges_all_starnet$node2, sep = "_")
edges_all_gtex$edge_name = paste(edges_all_gtex$node1, edges_all_gtex$node2,
    sep = "_")
fraction_tp_fp_fn <- function(edges_all, name, complete_name = "100_1",</pre>
    prob, clique_col, In_clique = 0) {
    tp = length(intersect(unique(edges_all$edge_name[edges_all$name ==
        complete name]), edges all$edge name[edges all$name ==
        name & edges_all$prob == prob & edges_all[, clique_col] ==
        In clique]))
    fp = length(setdiff(edges_all$edge_name[edges_all$name ==
        name & edges_all$prob == prob & edges_all[, clique_col] ==
        In clique], unique(edges all$edge name[edges all$name ==
        complete name])))
    fn = length(setdiff(unique(edges_all$edge_name[edges_all$name ==
        complete_name]), edges_all$edge_name[edges_all$name ==
        name & edges_all$prob == prob & edges_all[, clique_col] ==
        In_clique]))
   return(data.frame(TP = tp, FP = fp, FN = fn))
}
edges_all_starnet$name = paste(edges_all_starnet$data_type, edges_all_starnet$repRun,
    sep = " ")
edges_all_gtex$name = paste(edges_all_gtex$data_type, edges_all_gtex$repRun,
edges all starnet$prob = as.numeric(as.character(edges all starnet$postProb))
edges_all_gtex$prob = as.numeric(as.character(edges_all_gtex$postProb))
fig_2c_data = c()
for (prob in unique(edges_all_starnet$prob)) {
    # cat('----',prob,'----\n')
    for (subsampling in c(50, 80, 90, 100)) {
        for (rep in 1:5) {
            for (clique_col in c("inClique_0.01", "inClique_0.05",
                "inClique_0.1")) {
                gtex_NoClique = cbind(data.frame(prob = prob,
                  name = subsampling, run = rep, Data_source = "GTEx",
                  clique = clique_col, InClique = "No"), fraction_tp_fp_fn(edges_all_gtex,
                  paste(subsampling, rep, sep = "_"), "100_1",
                  prob, clique_col, 0))
                gtex YesClique = cbind(data.frame(prob = prob,
                  name = subsampling, run = rep, Data_source = "GTEx",
```

```
clique = clique_col, InClique = "Yes"), fraction_tp_fp_fn(edges_all_gtex,
                  paste(subsampling, rep, sep = "_"), "100_1",
                  prob, clique_col, 1))
                starnet_NoClique = cbind(data.frame(prob = prob,
                  name = subsampling, run = rep, Data_source = "STARNET",
                  clique = clique_col, InClique = "No"), fraction_tp_fp_fn(edges_all_starnet,
                  paste(subsampling, rep, sep = "_"), "100_1",
                  prob, clique col, 0))
                starnet_YesClique = cbind(data.frame(prob = prob,
                  name = subsampling, run = rep, Data_source = "STARNET",
                  clique = clique_col, InClique = "Yes"), fraction_tp_fp_fn(edges_all_starnet,
                  paste(subsampling, rep, sep = "_"), "100_1",
                  prob, clique_col, 1))
                fig_2c_data = rbind(fig_2c_data, gtex_NoClique,
                  gtex_YesClique, starnet_NoClique, starnet_YesClique)
           }
       }
   }
}
fig_2c_data$precision = fig_2c_data$TP/(fig_2c_data$TP + fig_2c_data$FP)
fig_2c_data$recall = fig_2c_data$TP/(fig_2c_data$TP + fig_2c_data$FN)
fig_2c_data = fig_2c_data[!is.na(fig_2c_data$precision), ]
fig_2c_summary = summarySE(fig_2c_data, "precision", groupvars = c("prob",
    "name", "Data_source", "clique", "InClique"))
fig_2c_summary$name = as.character(fig_2c_summary$name)
fig2c <- ggplot(fig_2c_summary, aes(prob)) + geom_line(aes(y = precision,
    colour = name)) + geom_ribbon(aes(ymin = precision - sd,
   ymax = precision + sd, fill = name), alpha = 0.6) + facet_wrap(~clique +
    Data_source + InClique, ncol = 4) + theme_bw() + scale_fill_manual(values = c("black",
    "cyan4", "magenta", "yellow4")) + scale_color_manual(values = c("black",
    "cyan4", "magenta", "yellow4")) + scale_x_continuous(breaks = seq(0.1,
   0.9, 0.4))
show(fig2c)
```

Figure 3: Simulation Data Precision / Recall

```
load("Simulation_edges_all_RIMBAnet.RData")
real_network = read.delim(paste(gitPath, "data/Simulation_true_network.txt",
    sep = ""), header = F)
colnames(real_network) = c("node1", "node2")

tpr <- function(edges, real_network) {
    edges_test = paste(edges$node1, edges$node2, sep = "_")
    edges_real = paste(real_network$node1, real_network$node2,
        sep = "_")
    return(length(intersect(edges_test, edges_real))/length(edges_test))
}</pre>
```

```
fdr <- function(edges, real_network) {</pre>
    edges_test = paste(edges$node1, edges$node2, sep = "_")
    edges_real = paste(real_network$node1, real_network$node2,
    return(length(setdiff(edges_test, edges_real))/length(edges_test))
}
recall <- function(edges, real network) {</pre>
    edges_test = paste(edges$node1, edges$node2, sep = "_")
    edges_real = paste(real_network$node1, real_network$node2,
        sep = " ")
    return(length(intersect(edges_test, edges_real))/length(edges_real))
}
tpr_res = data.frame()
fdr_res = data.frame()
recall_res = data.frame()
for (p in seq(0.1, 0.9, 0.1)) {
    for (name in c(10, 50, 80, 90, 100)) {
        for (rep in 1:5) {
            test_data = edges_all[edges_all$postProb == p & edges_all$data_type ==
                name & edges_all$repRun == rep, ]
            fdr_res = rbind(fdr_res, data.frame(name = name,
                repRun = rep, postProb = p, fdr = fdr(test_data,
                  real network)))
            tpr_res = rbind(tpr_res, data.frame(name = name,
                repRun = rep, postProb = p, tpr = tpr(test_data,
                  real_network)))
            recall_res = rbind(recall_res, data.frame(name = name,
                repRun = rep, postProb = p, recall = recall(test_data,
                  real network)))
        }
    }
}
library(ggplot2)
tpr_res$repRun = factor(tpr_res$repRun)
tpr_res2 = tpr_res[is.element(tpr_res$postProb, c(seq(0.1, 0.9,
    0.2))), ]
tpr_res2$name = tpr_res2$name * 10
tpr_sum = summarySE(tpr_res[is.element(tpr_res$postProb, c(seq(0.1,
    0.9, 0.4))), ], "tpr", groupvars = c("name", "postProb"))
tpr_sum$postProb2 = as.character(tpr_sum$postProb)
tpr_sum$name = tpr_sum$name * 10
fdr_sum = summarySE(fdr_res[is.element(fdr_res$postProb, c(seq(0.1,
    0.9, 0.4))), ], "fdr", groupvars = c("name", "postProb"))
fdr_res$repRun = factor(fdr_res$repRun)
fdr_res2 = fdr_res[is.element(fdr_res$postProb, c(seq(0.1, 0.9,
    0.2))), ]
```

```
fdr_res2$name = fdr_res2$name * 10
recall_sum = summarySE(recall_res[is.element(recall_res$postProb,
    c(seq(0.1, 0.9, 0.4))), ], "recall", groupvars = c("name",
    "postProb"))
recall_sum$postProb2 = as.character(recall_sum$postProb)
recall_sum$name = recall_sum$name * 10
## precision recall:
tpr_res$type = "Precision"
recall_res$type = "Recall"
colnames(tpr_res)[colnames(tpr_res) == "tpr"] = "value"
colnames(recall_res)[colnames(recall_res) == "recall"] = "value"
total = rbind(tpr_res, recall_res)
total_sum = summarySE(total, "value", groupvars = c("name", "postProb",
total_sum$postProb2 = as.character(total_sum$postProb)
total_sum$postProb_type = paste(total_sum$postProb2, total_sum$type,
    sep = " ")
total_sum$name = total_sum$name * 10
```

Figure 3:

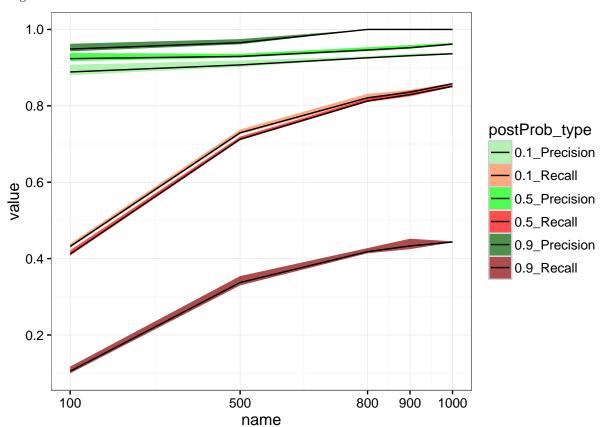


Figure 4: Reproducibility of Key Driver Nodes (KDs)

```
options(stringsAsFactors = F)
rm(list=ls())
library(ggplot2)
library("igraph")
library(plyr)
gitPath <- c("https://raw.githubusercontent.com/divara01/PSB2017_ReproducibilityOfBNs/")</pre>
## setting up the files for KD Calling --
load(paste(gitPath, "results/", "GTEx_edges_all_RIMBAnet_cliqueAdded.RData", sep = ""))
edges_all$type_name = paste(edges_all$data_type,edges_all$repRun,sep="_")
for(p in unique(edges_all$postProb)){
  for(d in unique(edges_all$type_name)){
    write.table(edges_all[edges_all$postProb == p & edges_all$type_name == d,
                          c('node1', 'node2')], sep = "\t", quote = F, row.names = F,
                col.names = F, file = paste("BN", d, p, "edges_GTEx.txt", sep="_"))
 }
}
## setting up the files for KD Calling --
load(paste(gitPath, "results/", "STARNET_edges_all_RIMBAnet_cliqueAdded.RData", sep = ""))
edges_all$type_name = paste(edges_all$data_type,edges_all$repRun,sep="_")
for(p in unique(edges_all$postProb)){
  for(d in unique(edges_all$type_name)){
    write.table(edges_all[edges_all$postProb == p & edges_all$type_name == d,
                          c('node1', 'node2')], sep = "\t", quote = F, row.names = F,
                col.names = F, file = paste("BN", d, p, "edges_STARNET.txt", sep = "_"))
 }
}
options(stringsAsFactors=F)
rm(list=ls())
args <- commandArgs(trailingOnly=TRUE) # disease = list of diseases split by ",", outputName
datasets <- c("GTEx", "STARNET")</pre>
for(datasetIndex in datasets)
  #Load network edge file
  load(paste(gitPath, "results/", datasetIndex, "edges_all_RIMBAnet_cliqueAdded.RData", sep = ""))
  Edge_File <- edges_all[, c(1, 2)]</pre>
  colnames(Edge_File) <- c("From", "To")</pre>
  #Create network in R
  #Determine connectivity between all genes in BN
  InverseLengthAllShortestPaths <- vector("list", 1)</pre>
  edgeDF_i <- Edge_File
```

```
g_i <- graph.data.frame(d = edgeDF_i, directed = TRUE)</pre>
  summedInverseLengthAllShortestPaths <- vector("list", length(V(g i)))</pre>
  for(node_i in 1:length(V(g_i))){
    SP_i <- get.all.shortest.paths(g_i, node_i, to = V(g_i), mode = "out")
    if(length(SP_i$res) > 0){
      #This tells you how many nodes between the starting node to the target node
      shortestPathLength i <- sapply(SP i$res, length)</pre>
      #Identifies the final target gene in the path
      target_i <- sapply(SP_i$res, function(x) tail(x,1))</pre>
      #number of shortest paths to the target
      spPerTarget <- split(x = shortestPathLength_i, f = target_i)</pre>
      #Calculates the score for each target gene from the source gene (1/#shortest paths)
      scorePerTarget <- sapply(spPerTarget, function(x) (1/unique(x)) * length(x))</pre>
      #Assigns a score of 0 to all genes in the subgraph
      op_i <- rep(0, length(V(g_i)))</pre>
      #This adds the score to the genes that were included in the shortest paths.
      #Remaining genes are still 0
      op_i[as.numeric(names(scorePerTarget))] <- as.numeric(scorePerTarget)</pre>
      summedInverseLengthAllShortestPaths[[node i]] <- op i</pre>
    }
  }
  #This converts the list of scores into a dataframe
  summedInverseLengthAllShortestPaths <- ldply(summedInverseLengthAllShortestPaths, rbind)</pre>
  #qene IDs are provided as row names
  rownames(summedInverseLengthAllShortestPaths) <- V(g_i)$name</pre>
  #gene IDs are provided as column names
  colnames(summedInverseLengthAllShortestPaths) <- V(g_i)$name</pre>
  totalScorePerGene <- rowSums(summedInverseLengthAllShortestPaths)</pre>
  quantileThr <- quantile(rowSums(summedInverseLengthAllShortestPaths), 0.95)
  KeyDrivers <- totalScorePerGene[which(totalScorePerGene >= quantileThr)]
  KD flag <- as.integer(names(totalScorePerGene) %in% names(KeyDrivers))</pre>
  perc.rank <- function(x, xo) length(x[x <= xo])/length(x)*100</pre>
  KDpercentile <- sapply(totalScorePerGene, function(x) perc.rank(totalScorePerGene, x))</pre>
  top_KDs = KDpercentile[KDpercentile >= 95]
  top_KDs = sort(top_KDs,decreasing=T)
  outputFN = paste(datasetIndex,"_KDs.txt",sep="")
  write.table(top_KDs,file = outputFN, sep="\t",quote=F, row.names=T, col.names=F)
}
options(stringsAsFactors=F)
rm(list=ls())
files = list.files(".","_STARNET_KDs.txt")
all kda = c()
```

```
for(file in files){
  x = read.delim(file,header=F)
  data_type = strsplit(file,"_")[[1]][2]
  repRun = strsplit(file, "_")[[1]][3]
  postProb = strsplit(file,"_")[[1]][4]
  all_kda = rbind(all_kda, data.frame(gene = x[,1], data_type = data_type, repRun = repRun,
                                      postProb = postProb))
}
load(paste(gitPath, "results/", "STARNET_edges_all_RIMBAnet_cliqueAdded.RData", sep = ""))
head(edges_all)
edges_all$KDflagNode1_redo = sapply(1:nrow(edges_all),function(x){
  is.element(edges_all$node1[x],all_kda$gene[all_kda$repRun == edges_all$repRun[x] &
                                               all_kda$data_type == edges_all$data_type[x] &
                                               all_kda$postProb == edges_all$postProb[x]])
})
table(edges_all$KDflagNode1,edges_all$KDflagNode1_redo)
save(edges_all, file = "STARNET_edges_all_RIMBAnet_cliqueAdded_KDAdded.RData")
## reading in the KD Results --
options(stringsAsFactors=F)
rm(list=ls())
files = list.files("."," GTEx KDs.txt")
all kda = c()
for(file in files){
 x = read.delim(file,header=F)
 data_type = strsplit(file,"_")[[1]][2]
 repRun = strsplit(file,"_")[[1]][3]
  postProb = strsplit(file,"_")[[1]][4]
  all_kda = rbind(all_kda, data.frame(gene = x[,1] , data_type = data_type,repRun = repRun,
                                      postProb = postProb))
}
load( "GTEx_edges_all_RIMBAnet_cliqueAdded.RData")
head(edges_all)
edges_all$KDflagNode1_redo = sapply(1:nrow(edges_all),function(x){
  is.element(edges_all$node1[x], all_kda$gene[all_kda$repRun == edges_all$repRun[x] &
                                                all_kda$data_type == edges_all$data_type[x] &
                                                all kda$postProb == edges all$postProb[x]])
})
table(edges_all$KDflagNode1,edges_all$KDflagNode1_redo)
save(edges_all, file = "GTEx_edges_all_RIMBAnet_cliqueAdded_KDAdded.RData")
```

Generate Node score for all nodes:

```
#Calculate Most Connected Node Conservation
datasets <- c("GTEx", "STARNET")
```

```
for(datasetIndex in 1:length(datasets)){
  message(paste(datasets[datasetIndex], "running..."))
  #loads in the edges all file
  load(paste(datasets[datasetIndex], " edges all RIMBAnet cliqueAdded KDAdded.RData", sep =""))
  allNodes <- unique(as.vector(unlist(edges_all[, c(1:2)])))</pre>
  totalRows <- length(unique(edges_all$postProb)) *</pre>
    length(unique(edges all$data type)) *
    length(unique(edges all$repRun)) *
    length(allNodes)
  postProb <- rep(sort(as.vector(unique(edges_all$postProb))),</pre>
                  totalRows/length(unique(edges_all$postProb)))
  subsets <- rep(as.vector(sapply(unique(edges_all$data_type), function(x)</pre>
    rep(x, length(unique(edges_all$postProb)) *
          length(unique(edges_all$repRun))))), length(allNodes))
  nodes <- as.vector((sapply(allNodes, function(x))</pre>
    rep(x, length(unique(edges all$data type)) * length(unique(edges all$repRun)) *
          length(unique(edges all$postProb))))))
  repRun <- rep(as.vector(sapply(as.vector(unique(edges all$repRun)), function(x)
    rep(x, length(unique(edges_all$postProb))))), length(allNodes) *
      length(unique(edges_all$data_type)))
  nodeTable <- as.data.frame(cbind(nodes, postProb, subsets, repRun))</pre>
  message(paste(datasets[datasetIndex], "evaluating each node's connections..."))
  for(i in 1:nrow(nodeTable)){
    node_i <- nodeTable$nodes[i]</pre>
    postProb_i <- nodeTable$postProb[i]</pre>
    subset_i <- nodeTable$subsets[i]</pre>
    repRun_i <- nodeTable$repRun[i]</pre>
    nodeTable$outDegree[i] <- length(which(edges_all$node1 == node_i &</pre>
                                               edges all$postProb == postProb i &
                                               edges all$data type == subset i &
                                               edges all$repRun == repRun i))
    nodeTable$inDegree[i] <- length(which(edges all$node2 == node i &</pre>
                                              edges_all$postProb == postProb_i &
                                              edges all$data type == subset i &
                                              edges_all$repRun == repRun_i))
    nodeTable$allDegree[i] <- nodeTable$outDegree[i] + nodeTable$inDegree[i]</pre>
    if(i %% 500 == 0){message(paste(nrow(nodeTable) - i, " remaining..."))}
  }
  assign(paste(datasets[datasetIndex], "allNodeScores", sep = "_"), nodeTable)
  write.table(nodeTable, file = paste(datasets[datasetIndex], "_allNodeScores", sep = ""),
              sep = "\t", row.names = FALSE, col.names = TRUE)
  message(paste(datasets[datasetIndex], "complete!"))
}
```

```
#Append values to the edges_all table
for(datasetIndex in 1:length(datasets)){
  message(paste(datasets[datasetIndex], "running..."))
  #loads in the edges all file
  load(paste(datasets[datasetIndex], "_edges_all_RIMBAnet_cliqueAdded_KDAdded.RData", sep =""))
  nodeTable <- read.delim(paste(datasets[datasetIndex], "_allNodeScores", sep = ""), sep = "\t",</pre>
                           header = TRUE)
  edges_all$outDegree <- NA
  edges_all$inDegree <- NA
  edges_all$allDegree <- NA
  for(i in 1:nrow(edges_all)){
    node1_i <- edges_all$node1[i]</pre>
    postProb_i <- edges_all$postProb[i]</pre>
    repRun_i <- edges_all$repRun[i]</pre>
    data_type_i <- edges_all$data_type[i]</pre>
    nodeTablePos <- which(nodeTable$nodes == node1_i &</pre>
                             nodeTable$postProb == postProb_i &
                             nodeTable$subsets == data_type_i &
                             nodeTable$repRun == repRun_i)
    edges_all$outDegree[i] <- nodeTable$outDegree[nodeTablePos]</pre>
    edges_all$inDegree[i] <- nodeTable$inDegree[nodeTablePos]</pre>
    edges_all$allDegree[i] <- nodeTable$allDegree[nodeTablePos]</pre>
    if(i %% 500 == 0){message(paste(nrow(edges_all) - i, " remaining..."))}
  }
  edgesAll_filename <- paste(dataset[datasetIndex],</pre>
                               "_edges_all_RIMBAnet_cliqueAdded_KDAdded_HubNodes.RData", sep = "")
  save(edges all,file=edgesAll filename)
  message(paste(datasets[datasetIndex], "Complete!"))
```

Figure 5: Hub Nodes Reproducibility Rate

```
options(stringsAsFactors = F)
rm(list = ls())
require(ggplot2)
require(plyr)
require(gridExtra)
require(grid)
library(ggplot2)

gtex = read.delim("GTEx_allNodeScores")
starnet = read.delim("STARNET_allNodeScores")
gtex = gtex[gtex$allDegree != 0, ]
starnet = starnet[starnet$allDegree != 0, ]
```

```
starnet_fn = "STARNET_edges_all_RIMBAnet_cliqueAdded_KDAdded_HubNodes.RData"
gtex_fn = "GTEx_edges_all_RIMBAnet_cliqueAdded_KDAdded_HubNodes.RData"
load(starnet fn)
edges_all_starnet = edges_all
rm(edges all)
load(gtex fn)
edges_all_gtex = edges_all
rm(edges_all)
edges_all_starnet$name = paste(edges_all_starnet$data_type, edges_all_starnet$repRun,
edges_all_gtex$name = paste(edges_all_gtex$data_type, edges_all_gtex$repRun,
    sep = " ")
edges_all_starnet = edges_all_starnet[edges_all_starnet$postProb !=
edges_all_gtex = edges_all_gtex[edges_all_gtex$postProb != 1,
   ]
gtex = gtex[gtex$postProb != 1, ]
starnet = starnet[starnet$postProb != 1, ]
## calculate the top connected out degree nodes:
gtex_top = data.frame()
starnet_top = data.frame()
for (p in unique(gtex$postProb)) {
   for (name in c(50, 80, 90, 100)) {
        for (rep in 1:5) {
            if (name == 100) {
                # for the complete networks, taking the same cutoff as the
                # 100_1 in all replicates because edge number is so similar
                # to avoid the issue of slight biase by different cutoffs.
                gtex_top = rbind(gtex_top, data.frame(postProb = p,
                  data_type = name, repRun = rep, outCutoff = quantile(gtex$outDegree[gtex$repRun ==
                    1 & gtex$postProb == p & gtex$subsets ==
                    name], 0.9)[[1]]))
                starnet_top = rbind(starnet_top, data.frame(postProb = p,
                  data_type = name, repRun = rep, outCutoff = quantile(starnet$outDegree[starnet$repRun
                    1 & starnet$postProb == p & starnet$subsets ==
                    name], 0.9)[[1]]))
            } else {
                gtex_top = rbind(gtex_top, data.frame(postProb = p,
                  data_type = name, repRun = rep, outCutoff = quantile(gtex$outDegree[gtex$repRun ==
                    rep & gtex$postProb == p & gtex$subsets ==
                    name], 0.9)[[1]]))
                starnet_top = rbind(starnet_top, data.frame(postProb = p,
                  data_type = name, repRun = rep, outCutoff = quantile(starnet$outDegree[starnet$repRun
                    rep & starnet$postProb == p & starnet$subsets ==
                    name], 0.9)[[1]]))
           }
       }
```

```
gtex$name = paste(gtex$subsets, gtex$repRun, gtex$postProb, sep = "_")
starnet$name = paste(starnet$subsets, starnet$repRun, starnet$postProb,
starnet_top$name = paste(starnet_top$data_type, starnet_top$repRun,
    starnet top$postProb, sep = " ")
gtex_top$name = paste(gtex_top$data_type, gtex_top$repRun, gtex_top$postProb,
    sep = " ")
## for each node1 assign if it is in the top connected node:
edges_all_starnet$name = paste(edges_all_starnet$data_type, edges_all_starnet$repRun,
    edges_all_starnet$postProb, sep = "_")
edges_all_gtex$name = paste(edges_all_gtex$data_type, edges_all_gtex$repRun,
    edges_all_starnet$postProb, sep = "_")
starnet_top1 = starnet_top$outCutoff
names(starnet_top1) = starnet_top$name
edges_all_starnet$TopConnNode1 = sapply(1:nrow(edges_all_starnet),
    function(x) {
        is.element(edges_all_starnet$node1[x], starnet$nodes[starnet$name ==
            edges_all_starnet$name[x] & starnet$outDegree >=
            starnet_top1[edges_all_starnet$name[x]]])
   })
gtex_top1 = gtex_top$outCutoff
names(gtex_top1) = gtex_top$name
edges_all_gtex$TopConnNode1 = sapply(1:nrow(edges_all_gtex),
   function(x) {
        is.element(edges_all_gtex$node1[x], gtex$nodes[gtex$name ==
            edges_all_gtex$name[x] & gtex$outDegree >= gtex_top1[edges_all_gtex$name[x]]])
   })
kda = rbind(data.frame(unique(edges_all_gtex[, c("node1", "postProb",
    "repRun", "data_type", "TopConnNode1")]), Data_source = "GTEx"),
    data.frame(unique(edges_all_starnet[, c("node1", "postProb",
        "repRun", "data_type", "TopConnNode1")]), Data_source = "STARNET"))
kda$name = paste(kda$Data_source, kda$data_type, kda$repRun,
    sep = "_")
kda$postProb = kda$postProb
kda = kda[kda$TopConnNode1 & kda$postProb != 1, ]
getFractionOverlap = function(kda, A_name, B_name, prob, fixed = TRUE) {
    if (A_name == B_name) {
        A_size = strsplit(A_name, "_")[[1]][2]
        B_name = setdiff(unique(kda$name[kda$data_type != A_size]),
            A_name)
   }
    if (fixed) {
        AandB = length(intersect(kda$node1[kda$name == A_name &
            kda$postProb == prob], kda$node1[is.element(kda$name,
            B_name) & kda$postProb == prob]))
```

```
} else {
        AandB = length(intersect(kda$node1[kda$name == A_name &
            kda$postProb == prob], kda$node1[is.element(kda$name,
    a = length(unique(kda$node1[kda$name == A_name & kda$postProb ==
        prob]))
   return(AandB/a)
}
kda_fractions = data.frame()
subsamp = c(50, 80, 90, 100)
for (p in unique(kda$postProb)) {
   for (sub in unique(kda$data_type)) {
        for (k in unique(kda$repRun)) {
            for (fixed in c(TRUE, FALSE)) {
                for (dataSet in unique(kda$Data_source)) {
                  # cat('on ', dataSet, sub, k, fixed, p, ' n')
                  if (sub == 100 \& k == "1") {
                    break
                  kda_fractions = rbind(kda_fractions, data.frame(Data_source = dataSet,
                    data_type = sub, repRun = k, TP_fraction = getFractionOverlap(kda,
                      paste(dataSet, sub, k, sep = "_"), paste(dataSet,
                        100, 1, sep = "_"), p, fixed), fixed = fixed,
                    postProb = p, splitBy = "No Split"))
            }
       }
   }
}
kda_fractions$repRun = factor(kda_fractions$repRun)
kda_fractions$fixed = factor(kda_fractions$fixed, levels = c("TRUE",
    "FALSE"))
kda_fractions$postProb = factor(kda_fractions$postProb)
kda_fractions$data_type = as.character(kda_fractions$data_type)
kda_fractions2 = kda_fractions[is.element(kda_fractions$postProb,
    c(seq(0.1, 0.5, 0.1))), ]
kda_summary <- summarySE(kda_fractions, "TP_fraction", groupvars = c("fixed",
    "postProb", "data_type", "Data_source"))
kda_summary$data_type = as.character(kda_summary$data_type)
kda_summary$postProb = as.numeric(as.character(kda_summary$postProb))
fig5 <- ggplot(kda_summary, aes(postProb)) + geom_line(aes(y = TP_fraction,
    colour = data_type), size = 1) + geom_ribbon(aes(ymin = TP_fraction -
    sd, ymax = TP_fraction + sd, fill = data_type), alpha = 0.6) +
    scale_color_manual(values = c("black", "cyan4", "magenta",
        "yellow4")) + scale_fill_manual(values = c("black", "cyan4",
```

```
"magenta", "yellow4")) + facet_wrap(~fixed + Data_source,
ncol = 2) + theme_bw() + scale_x_continuous(breaks = seq(0.1,
0.9, 0.4))
show(fig5)
```

Table 2: Compare 5 Replicates of 100% Networks

```
options(stringsAsFactors = F)
rm(list = ls())
load(file = "STARNET edges all RIMBAnet cliqueAdded KDAdded HubNodes.RData")
edges_all_starnet = edges_all
edges_all_starnet$data_source = "STARNET"
rm(edges_all)
load(file = "GTEx_edges_all_RIMBAnet_cliqueAdded_KDAdded_HubNodes.RData")
edges_all_gtex = edges_all
edges_all_gtex$data_source = "GTEx"
rm(edges_all)
edges_all = rbind(edges_all_starnet, edges_all_gtex)
edges_all$unique_name = paste(edges_all$data_source, edges_all$data_type,
    edges_all$repRun, sep = "_")
edges_all$edge_name = paste(edges_all$node1, edges_all$node2,
   sep = "_")
edges all = edges all[edges all$data type == 100, ]
edges_all = edges_all[edges_all$postProb != 1, ]
testsToRun = combn(1:5, 2)
ab_ov_a = function(edges_all, a_name, b_name, prob1 = p, prob2 = p2) {
    ab = length(intersect(edges_all$edge_name[edges_all$unique_name ==
        a_name & edges_all$postProb == prob1], edges_all$edge_name[edges_all$unique_name ==
        b_name & edges_all$postProb == prob2]))
   a = sum(edges_all$unique_name == a_name & edges_all$postProb ==
        prob1)
   return(ab/a)
}
comp_res = data.frame()
for (i in 1:ncol(testsToRun)) {
    cat(i, "\n")
   for (p in unique(edges_all$postProb)) {
        comp_res = rbind(comp_res, data.frame(A_name = testsToRun[1,
            i], B name = testsToRun[2, i], data source = "GTEx",
            ab_ov_a = ab_ov_a(edges_all, paste("GTEx", 100, testsToRun[1,
                i], sep = "_"), paste("GTEx", 100, testsToRun[2,
                i], sep = "_"), prob1 = p, prob2 = p), postProb = p))
        comp_res = rbind(comp_res, data.frame(A_name = testsToRun[2,
            i], B_name = testsToRun[1, i], data_source = "GTEx",
```

```
ab_ov_a = ab_ov_a(edges_all, paste("GTEx", 100, testsToRun[2,
                i], sep = "_"), paste("GTEx", 100, testsToRun[1,
                i], sep = "_"), prob1 = p, prob2 = p), postProb = p))
        comp_res = rbind(comp_res, data.frame(A_name = testsToRun[1,
            i], B_name = testsToRun[2, i], data_source = "STARNET",
            ab_ov_a = ab_ov_a(edges_all, paste("STARNET", 100,
                testsToRun[1, i], sep = "_"), paste("STARNET",
                100, testsToRun[2, i], sep = "_"), prob1 = p,
                prob2 = p), postProb = p))
        comp_res = rbind(comp_res, data.frame(A_name = testsToRun[2,
            i], B_name = testsToRun[1, i], data_source = "STARNET",
            ab_ov_a = ab_ov_a(edges_all, paste("STARNET", 100,
                testsToRun[2, i], sep = "_"), paste("STARNET",
                100, testsToRun[1, i], sep = "_"), prob1 = p,
                prob2 = p), postProb = p))
    }
}
tab_out = data.frame(postProb = seq(0.1, 0.9, 0.1), GTEx_mean = NA,
    GTEx_sd = NA, STARNET_mean = NA, STARNET_sd = NA)
tab_out$GTEx_mean = sapply(tab_out$postProb, function(x) {
    mean(comp_res$ab_ov_a[comp_res$data_source == "GTEx" & comp_res$postProb ==
        x])
})
tab_out$GTEx_sd = sapply(tab_out$postProb, function(x) {
    sd(comp_res$ab_ov_a[comp_res$data_source == "GTEx" & comp_res$postProb ==
        ([x
})
tab_out$STARNET_mean = sapply(tab_out$postProb, function(x) {
    mean(comp_res$ab_ov_a[comp_res$data_source == "STARNET" &
        comp_res$postProb == x])
})
tab_out$STARNET_sd = sapply(tab_out$postProb, function(x) {
    sd(comp_res$ab_ov_a[comp_res$data_source == "STARNET" & comp_res$postProb ==
        ([x
})
write.table(tab_out, sep = "\t", row.names = F, col.names = T,
    file = "Updated_Table3.txt", quote = F)
```

Table 3: Jaccard Index

```
get_jacard <- function(edges_all, name1, name2, prob) {
   aANDb = length(intersect(edges_all$edge_name[edges_all$postProb ==
        prob & edges_all$name == name1], edges_all$edge_name[edges_all$postProb ==
        prob & edges_all$name == name2]))
   a = sum(edges_all$postProb == prob & edges_all$name == name1)</pre>
```

```
b = sum(edges_all$postProb == prob & edges_all$name == name2)
    jacard = aANDb/(a + b - aANDb)
    return(jacard)
}
edges_all_gtex$name = paste(edges_all_gtex$data_type, edges_all_gtex$repRun,
edges_all_starnet$name = paste(edges_all_starnet$data_type, edges_all_starnet$repRun,
    sep = "_")
edges_all_gtex$edge_name = paste(edges_all_gtex$node1, edges_all_gtex$node2,
    sep = " ")
edges_all_starnet$edge_name = paste(edges_all_starnet$node1,
    edges_all_starnet$node2, sep = "_")
jaccard_values = data.frame()
combos = combn(5, 2)
for (subsamp in c(50, 80, 90, 100)) {
    # cat('on subsampling', subsamp, '\n')
    for (p in seq(0.1, 0.9, 0.1)) {
        # cat('on prob',p,'\n')
        for (k in 1:ncol(combos)) {
            jaccard_values = rbind(jaccard_values, data.frame(name1 = paste(subsamp,
                combos[1, k], sep = "_"), name2 = paste(subsamp,
                combos[2, k], sep = "_"), prob = p, jaccard = get_jacard(edges_all_gtex,
                paste(subsamp, combos[1, k], sep = "_"), paste(subsamp,
                  combos[2, k], sep = "_"), p), dataSource = "GTEx"))
            jaccard_values = rbind(jaccard_values, data.frame(name1 = paste(subsamp,
                combos[1, k], sep = "_"), name2 = paste(subsamp,
                combos[2, k], sep = "_"), prob = p, jaccard = get_jacard(edges_all_starnet,
                paste(subsamp, combos[1, k], sep = "_"), paste(subsamp,
                  combos[2, k], sep = "_"), p), dataSource = "STARNET"))
            if (subsamp != 100) {
                jaccard_values = rbind(jaccard_values, data.frame(name1 = paste(subsamp,
                  combos[1, k], sep = "_"), name2 = "100_1",
                  prob = p, jaccard = get_jacard(edges_all gtex,
                    paste(subsamp, combos[1, k], sep = "_"),
                    "100_1", p), dataSource = "GTEx"))
                jaccard_values = rbind(jaccard_values, data.frame(name1 = paste(subsamp,
                  combos[1, k], sep = "_"), name2 = "100_1",
                  prob = p, jaccard = get_jacard(edges_all_starnet,
                    paste(subsamp, combos[1, k], sep = "_"),
                    "100 1", p), dataSource = "STARNET"))
            }
       }
   }
}
jaccard_values_output = jaccard_values[is.element(jaccard_values$prob,
    seq(0.1, 0.9, 0.4)), ]
jaccard_values_output$type1 = sapply(jaccard_values_output$name1,
    function(x) {
        strsplit(x, "_")[[1]][1]
```

```
})
jaccard_values_output$type2 = sapply(jaccard_values_output$name2,
    function(x) {
        strsplit(x, "_")[[1]][1]
    })

jaccard_table = summarySE(jaccard_values_output, "jaccard", groupvars = c("dataSource",
        "type1", "type2", "prob"))
print(jaccard_table[, c("dataSource", "type1", "type2", "prob",
        "N", "jaccard", "sd")])
```

Table 4: Edge Counts

```
load("Simulation edges all RIMBAnet.RData")
edges_all_sim = edges_all
rm(edges all)
edges_all_sim$type_name = paste(edges_all_sim$data_type, edges_all_sim$repRun,
edges_all_sim$edge_name = paste(edges_all_sim$node1, edges_all_sim$node2,
    sep = " ")
load("GTEx_edges_all_RIMBAnet_cliqueAdded_KDAdded_HubNodes.RData")
edges_all_gtex = edges_all
edges_all_gtex$type_name = paste(edges_all_gtex$data_type, edges_all_gtex$repRun,
    sep = " ")
edges_all_gtex$edge_name = paste(edges_all_gtex$node1, edges_all_gtex$node2,
    sep = "_")
rm(edges_all)
load("STARNET_edges_all_RIMBAnet_cliqueAdded_KDAdded_HubNodes.RData")
edges all starnet = edges all
rm(edges_all)
edges_all_starnet$type_name = paste(edges_all_starnet$data_type,
    edges_all_starnet$repRun, sep = "_")
edges_all_starnet$edge_name = paste(edges_all_starnet$node1,
    edges_all_starnet$node2, sep = "_")
tab4_starnet = table(edges_all_starnet$type_name[is.element(edges_all_starnet$postProb,
    seq(0.1, 0.9, 0.4))], edges_all_starnet$postProb[is.element(edges_all_starnet$postProb,
    seq(0.1, 0.9, 0.4))])
colnames(tab4_starnet) = paste("STARNET", colnames(tab4_starnet))
edges_all_gtex$postProb = as.numeric(as.character(edges_all_gtex$postProb))
tab4_gtex = table(edges_all_gtex$type_name[is.element(edges_all_gtex$postProb,
    seq(0.1, 0.9, 0.4))], edges_all_gtex$postProb[is.element(edges_all_gtex$postProb,
    seq(0.1, 0.9, 0.4))])
colnames(tab4 gtex) = paste("GTEx", colnames(tab4 gtex))
edges_all_sim$postProb = as.numeric(as.character(edges_all_sim$postProb))
tab4_sim = table(edges_all_sim$type_name[is.element(edges_all_sim$postProb,
    seq(0.1, 0.9, 0.4))], edges_all_sim$postProb[is.element(edges_all_sim$postProb,
    seq(0.1, 0.9, 0.4))])
```

```
colnames(tab4_sim) = paste("Sim", colnames(tab4_sim))

tab4_starnet = rbind(matrix(NA, nrow = 5, ncol = 3), tab4_starnet)
tab4_gtex = rbind(matrix(NA, nrow = 5, ncol = 3), tab4_gtex)
rownames(tab4_starnet)[1:5] = rownames(tab4_sim)[1:5]
rownames(tab4_gtex)[1:5] = rownames(tab4_sim)[1:5]
tab4 = cbind(tab4_starnet, tab4_gtex, tab4_sim)
tab4_melt = melt(tab4)
tab4_melt$X1 = as.character(tab4_melt$X1)
tab4_melt$subsampling = sapply(tab4_melt$X1, function(x) {
    strsplit(x, "_")[[1]][1]
})
table4 <- summarySE(tab4_melt, "value", groupvars = c("subsampling",
    "X2"))
print(table4[!is.na(table4$value), c("subsampling", "X2", "N",
    "value", "sd")])</pre>
```