

# T-Cell Network

*Example for GeneNet 1.2.13 (August 2015) or later*

This note reproduces the “T-Cell” network example from R. Opgen-Rhein and K. Strimmer. 2006a. *Using regularized dynamic correlation to infer gene dependency networks from time-series microarray data*. Proceedings of WCSB 2006 (June 12-13, 2006, Tampere, Finland) and R. Opgen-Rhein and K. Strimmer. 2006b. *Inferring gene dependency networks from genomic longitudinal data: a functional data approach*. REVSTAT 4:53-65. (<http://www.ine.pt/revstat/pdf/rs060103.pdf>)

## Load GeneNet library

```
library("GeneNet")
```

```
## Loading required package: corpcor  
## Loading required package: longitudinal  
## Loading required package: fdrtool
```

get T cell data

```
data(tcell)  
tc44 = combine.longitudinal(tcell.10, tcell.34)
```

## Estimate partial correlations

```
pc1 = ggm.estimate.pcor(tc44, lambda=0) # static, no shrinkage
```

```
## Specified shrinkage intensity lambda (correlation matrix): 0
```

```
pc2 = ggm.estimate.pcor(tc44, method="dynamic", lambda=0) # dynamic, no shrinkage
```

```
## Specified shrinkage intensity lambda (correlation matrix): 0
```

```
pc3 = ggm.estimate.pcor(tc44) # static, with shrinkage
```

```
## Estimating optimal shrinkage intensity lambda (correlation matrix): 0.0209
```

```
pc4 = ggm.estimate.pcor(tc44, method="dynamic") # dynamic, with shrinkage
```

```
## Estimating optimal shrinkage intensity lambda (correlation matrix): 0.032
```

## Find significant edges

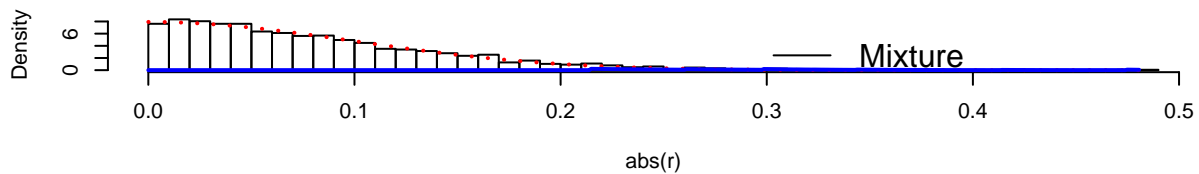
We use as selection criterion local fdr  $\leq 0.2$

static, no shrinkage

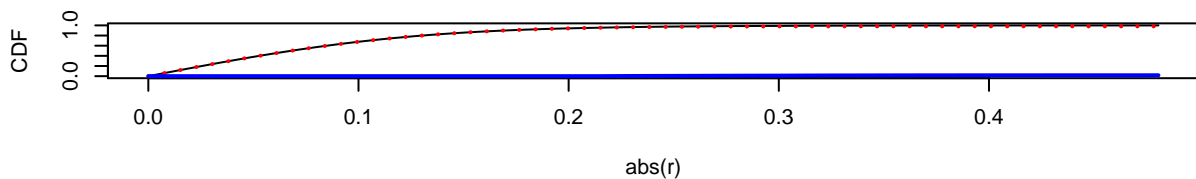
```
t1.edges = network.test.edges(pc1)
```

```
## Estimate (local) false discovery rates (partial correlations):  
## Step 1... determine cutoff point  
## Step 2... estimate parameters of null distribution and eta0  
## Step 3... compute p-values and estimate empirical PDF/CDF  
## Step 4... compute q-values and local fdr  
## Step 5... prepare for plotting
```

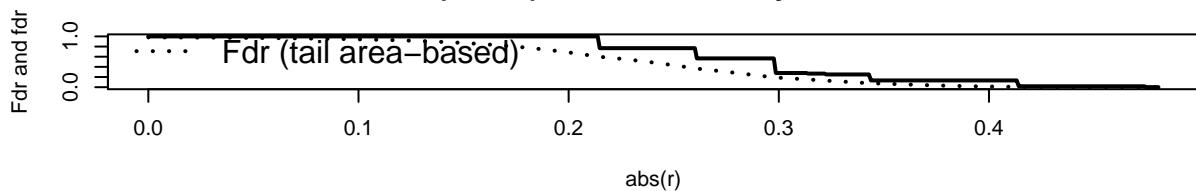
### Type of Statistic: Correlation (kappa = 104, eta0 = 0.9815)



### Density (first row) and Distribution Function (second row)



### (Local) False Discovery Rate



```
t1.net = extract.network(t1.edges) # prob > 0.8
```

```
##  
## Significant edges: 6  
## Corresponding to 0.36 % of possible edges
```

```
t1.net
```

```
##      pcor node1 node2      pval      qval      prob  
## 1  0.4806257   11   28 2.124826e-07 0.0002588653 0.9997411  
## 2  0.4744295   40   56 3.191076e-07 0.0002588653 0.9821251
```

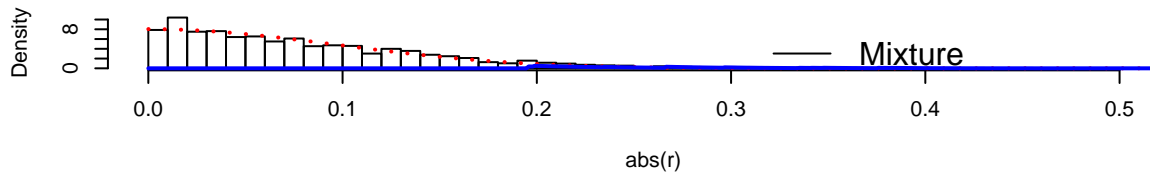
```
## 3 -0.4140609    31    45 1.133647e-05 0.0061308857 0.8673551
## 4  0.3635057    33    48 1.385462e-04 0.0493380622 0.8673551
## 5  0.3538857    23    28 2.132978e-04 0.0632622165 0.8673551
## 6  0.3444279    46    49 3.217399e-04 0.0768025473 0.8673551
```

dynamic, no shrinkage

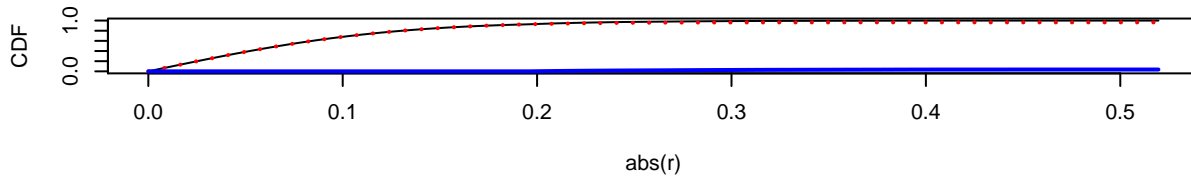
```
t2.edges = network.test.edges(pc2)
```

```
## Estimate (local) false discovery rates (partial correlations):
## Step 1... determine cutoff point
## Step 2... estimate parameters of null distribution and eta0
## Step 3... compute p-values and estimate empirical PDF/CDF
## Step 4... compute q-values and local fdr
## Step 5... prepare for plotting
```

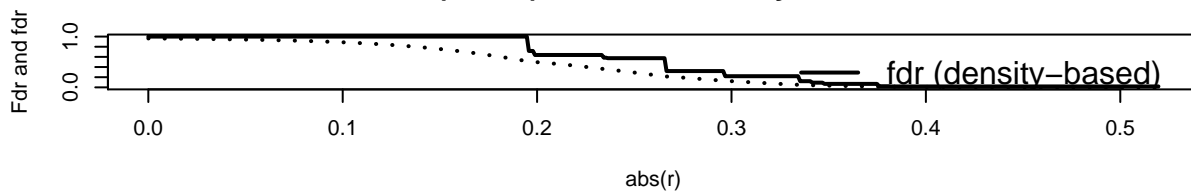
### Type of Statistic: Correlation (kappa = 110.7, eta0 = 0.964)



### Density (first row) and Distribution Function (second row)



### (Local) False Discovery Rate



```
t2.net = extract.network(t2.edges) # prob > 0.8
```

```
##
## Significant edges: 9
## Corresponding to 0.54 % of possible edges
```

```
t2.net
```

```
##          pcor node1 node2          pval          qval          prob
```

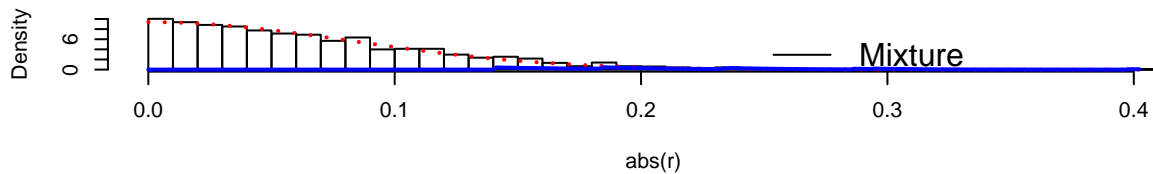
```
## 1 0.5196235 11 28 4.547772e-09 7.246755e-06 0.9821314
## 2 0.3971802 31 33 1.490293e-05 1.019816e-02 0.9821314
## 3 0.3888354 40 56 2.324970e-05 1.205612e-02 0.9821314
## 4 0.3817250 18 44 3.364533e-05 1.340324e-02 0.9778560
## 5 0.3749210 23 28 4.754204e-05 1.515140e-02 0.9318103
## 6 -0.3543569 5 26 1.291406e-04 2.979406e-02 0.9318103
## 7 0.3503019 23 32 1.560326e-04 3.299615e-02 0.9318103
## 8 0.3477015 46 49 1.759213e-04 3.504075e-02 0.9079267
## 9 0.3414539 30 40 2.337028e-04 4.137770e-02 0.8789744
```

static, with shrinkage

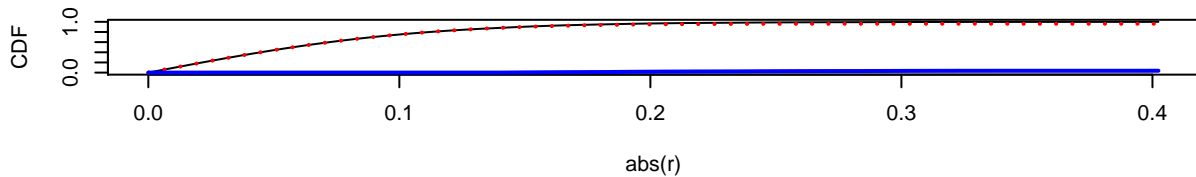
```
t3.edges = network.test.edges(pc3)
```

```
## Estimate (local) false discovery rates (partial correlations):
## Step 1... determine cutoff point
## Step 2... estimate parameters of null distribution and eta0
## Step 3... compute p-values and estimate empirical PDF/CDF
## Step 4... compute q-values and local fdr
## Step 5... prepare for plotting
```

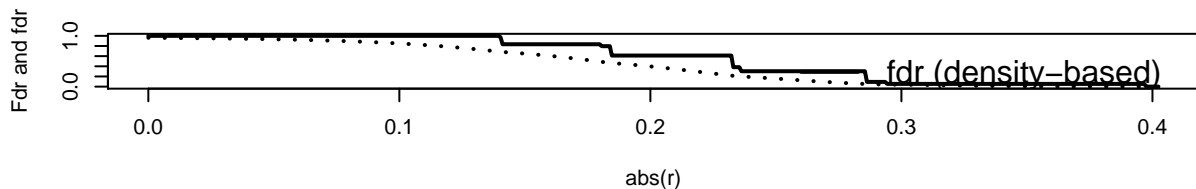
### Type of Statistic: Correlation (kappa = 151, eta0 = 0.9625)



### Density (first row) and Distribution Function (second row)



### (Local) False Discovery Rate



```
t3.net = extract.network(t3.edges) # prob > 0.8
```

```
##
## Significant edges: 10
## Corresponding to 0.6 % of possible edges
```

```
t3.net
```

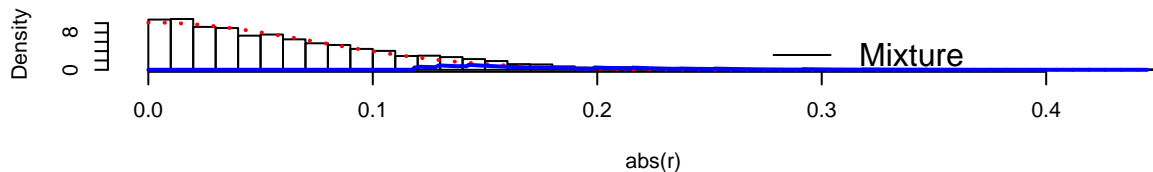
```
##          pcor node1 node2          pval          qval          prob
## 1  0.4023842   40   56 2.762444e-07 0.0003307888 0.9996692
## 2  0.3968325   11   28 4.158062e-07 0.0003307888 0.9463881
## 3  0.3164111   46   49 7.146148e-05 0.0276746523 0.9463881
## 4  0.3128258   23   28 8.710848e-05 0.0303085469 0.9463881
## 5  0.3093519   33   48 1.052786e-04 0.0327666910 0.9463881
## 6 -0.2961398   31   45 2.118707e-04 0.0407350046 0.9463881
## 7  0.2959314   31   33 2.141644e-04 0.0408400640 0.9463881
## 8  0.2954847    6   45 2.191583e-04 0.0410629683 0.9463881
## 9  0.2940215   29   36 2.362839e-04 0.0417716291 0.9070067
## 10 0.2892489    5   30 3.011663e-04 0.0473958579 0.9070067
```

dynamic, with shrinkage

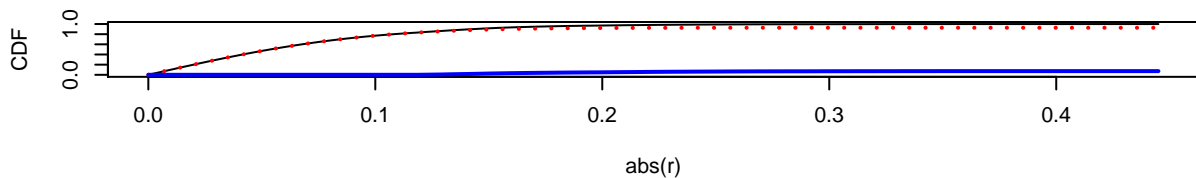
```
t4.edges = network.test.edges(pc4)
```

```
## Estimate (local) false discovery rates (partial correlations):
## Step 1... determine cutoff point
## Step 2... estimate parameters of null distribution and eta0
## Step 3... compute p-values and estimate empirical PDF/CDF
## Step 4... compute q-values and local fdr
## Step 5... prepare for plotting
```

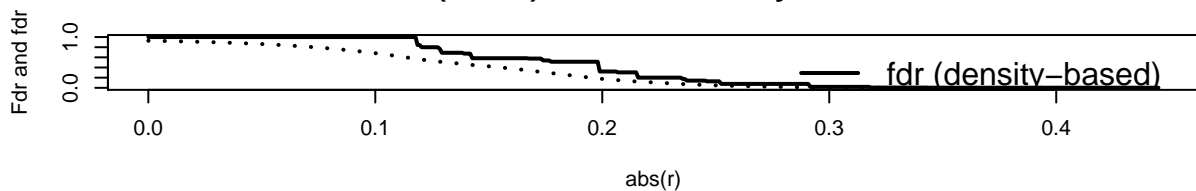
### Type of Statistic: Correlation ( $\kappa = 189.7$ , $\eta_0 = 0.9268$ )



### Density (first row) and Distribution Function (second row)



### (Local) False Discovery Rate



```
t4.net = extract.network(t4.edges) # prob > 0.8
```

```
##  
## Significant edges: 21  
## Corresponding to 1.27 % of possible edges
```

```
t4.net
```

```
##      pcor node1 node2      pval      qval      prob  
## 1  0.4450243   11   28 1.151954e-10 1.764880e-07 0.9980860  
## 2  0.3425852   18   44 1.249379e-06 9.570710e-04 0.9951212  
## 3  0.3217422   23   28 5.750408e-06 2.580987e-03 0.9951212  
## 4  0.3177284   40   56 7.618262e-06 2.917936e-03 0.9811075  
## 5  0.2987142   23   32 2.735755e-05 7.483611e-03 0.9811075  
## 6  0.2923531   46   49 4.114877e-05 9.382601e-03 0.9811075  
## 7  0.2910758   31   33 4.461217e-05 9.764167e-03 0.9231219  
## 8  0.2784745   37   51 9.700874e-05 1.847602e-02 0.9231219  
## 9  0.2670698    5   30 1.898769e-04 2.939947e-02 0.9231219  
## 10 -0.2660534   32   41 2.013014e-04 3.046735e-02 0.9231219  
## 11  0.2625146    4   55 2.462677e-04 3.424178e-02 0.9231219  
## 12  0.2555715   46   57 3.628202e-04 4.166472e-02 0.9231219  
## 13 -0.2553619    5   26 3.670299e-04 4.188476e-02 0.9231219  
## 14  0.2528018   30   40 4.222203e-04 4.453450e-02 0.9231219  
## 15  0.2517914   25   42 4.460449e-04 4.555827e-02 0.8718166  
## 16  0.2462722    3   16 5.996532e-04 5.456852e-02 0.8718166  
## 17  0.2458450   10   16 6.133780e-04 5.527887e-02 0.8581086  
## 18  0.2386917   38   58 8.907959e-04 6.825390e-02 0.8581086  
## 19  0.2382936    6   45 9.091928e-04 6.897824e-02 0.8581086  
## 20  0.2381243   49   56 9.171238e-04 6.928613e-02 0.8581086  
## 21 -0.2367511   40   45 9.838335e-04 7.177649e-02 0.8249718
```

## Produce plots using graph and Rgraphviz

```
library("Rgraphviz")
```

```
## Loading required package: graph  
## Loading required package: grid
```

```
node.labels = colnames(tc44)  
gr1 = network.make.graph( t1.net, node.labels)  
gr2 = network.make.graph( t2.net, node.labels)  
gr3 = network.make.graph( t3.net, node.labels)  
gr4 = network.make.graph( t4.net, node.labels)  
  
globalAttrs = list()  
globalAttrs$edge <- list(color = "blue", lty = "solid", lwd = 1, arrowsize=1)  
globalAttrs$node <- list(fillcolor = "lightyellow", shape = "ellipse", fixedsize = FALSE)  
  
edgeAttrs1 <- list()
```

```

edgeAttrs1$dir = edge.info(gr1)$dir
edgeAttrs2 <- list()
edgeAttrs2$dir = edge.info(gr2)$dir
edgeAttrs3 <- list()
edgeAttrs3$dir = edge.info(gr3)$dir
edgeAttrs4 <- list()
edgeAttrs4$dir = edge.info(gr4)$dir

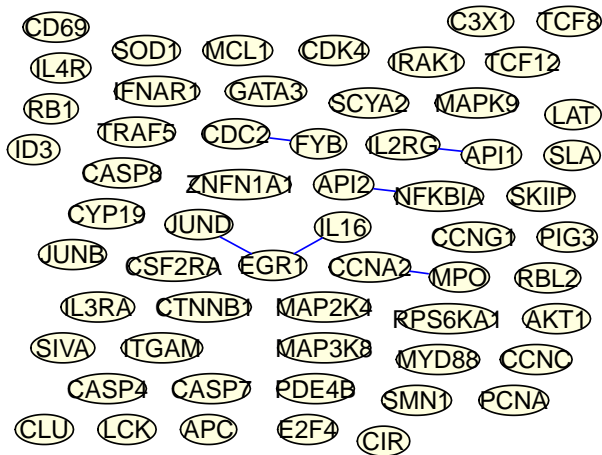
```

```

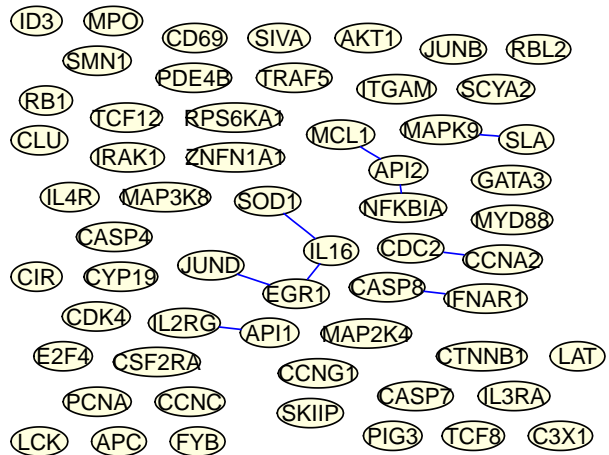
par(mfrow=c(2,2))
plot(gr1, main="Static, no shrinkage", attrs = globalAttrs, edgeAttrs = edgeAttrs1, "fdp")
plot(gr2, main="Dynamic, no shrinkage", attrs = globalAttrs, edgeAttrs = edgeAttrs2, "fdp")
plot(gr3, main="Static, with shrinkage", attrs = globalAttrs, edgeAttrs = edgeAttrs3, "fdp")
plot(gr4, main="Dynamic, with shrinkage", attrs = globalAttrs, edgeAttrs = edgeAttrs4, "fdp")

```

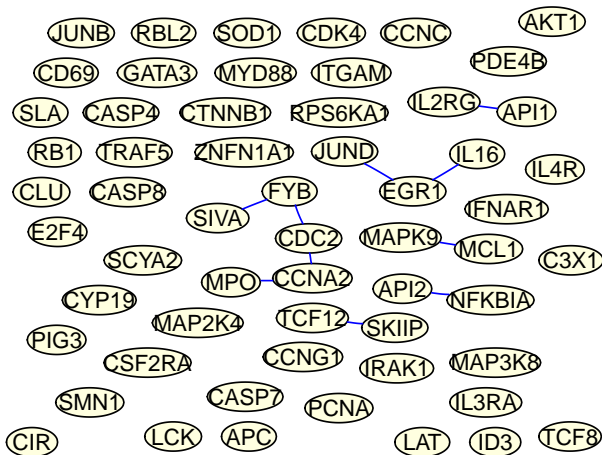
Static, no shrinkage



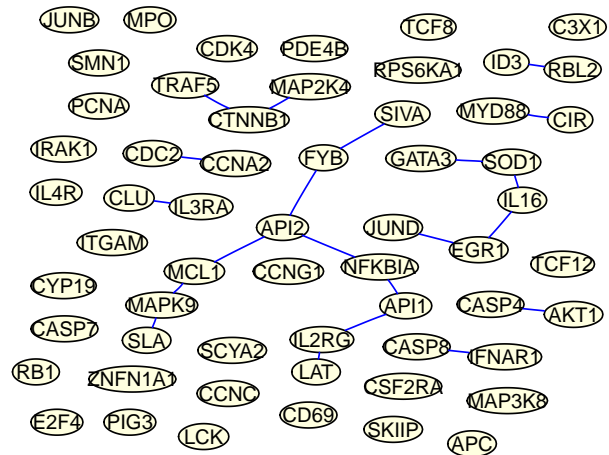
Dynamic, no shrinkage



Static, with shrinkage



Dynamic, with shrinkage



```

par(mfrow=c(1,1))

```