```
flashtpx: Simulation Run 1
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```

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Overview

This is the first simulation run of *flashtpx* without the non-negative constraint. The main idea is to try and replicate the results for the FitGoM() in CountClust or topics() model in the maptpx package due to Matt Taddy.

Without the non-negative constraint, *flashtpx* is basically applying *flash* with the covariance matrix for the data estimated from the GoM or topic model fitting on the counts data. Here we apply *flashtpx* on the counts data generated from a chosen simulation design and then interpret the results and compare the results to the PMA model fitting.

Simulation Design

We load the packages and the functions we need to perform the model.

```
library(ashr)
library(irlba)
library(PMA)
source("../R/flash.R")
```

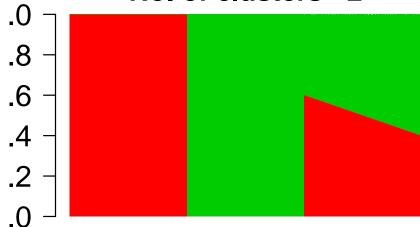
Next we determine the omega matrix determined in batches of 200, the first 200 batches of samples coming from one cluster completely, the next 200 coming from a second cluster and the final 200 being proportionally assigned to the two clusters. (Check barplot for more clarity).

```
n.out <- 200
omega_sim <- rbind(cbind(rep(1, n.out), rep(0,</pre>
    n.out)), cbind(rep(0, n.out), rep(1, n.out)),
    cbind(seq(0.6, 0.4, length.out = n.out), 1 -
        seq(0.6, 0.4, length.out = n.out)))
dim(omega_sim)
```

[1] 600

```
K <- dim(omega_sim)[2]</pre>
par(mar = c(2, 2, 2, 2))
barplot(t(omega\_sim), col = 2:(K + 1), axisnames = F,
    space = 0, border = NA, main = paste("No. of clusters=",
        K), las = 1, ylim = c(0, 1), cex.axis = 1.5,
    cex.main = 1.4)
```

No. of clusters= 2



So we have two clusters. How do these clusters look? Assume we have 100 genes.

```
freq <- rbind(c(0.1, 0.2, rep(0.7/98, 98)), c(rep(0.7/98,
    98), 0.1, 0.2))
str(freq)
```

```
## num [1:2, 1:100] 0.1 0.00714 0.2 0.00714 0.00714 ...
```

So the first cluster has high expression at first 2 genes and low expression at the other 98 genes.

```
counts <- t(do.call(cbind, lapply(1:dim(omega_sim)[1],</pre>
    function(x) rmultinom(1, 1000, prob = omega_sim[x,
        ] %*% freq))))
dim(counts)
```

[1] 600 100

We next fit a standard topic model with K = 2.

```
topic.fit <- maptpx::topics(counts, K = 2)</pre>
##
## Estimating on a 600 document collection.
## Fitting the 2 topic model.
## log posterior increase: 5599.4, 34.2, 0.3, done.
omega <- topic.fit$omega
theta <- topic.fit$theta
```

Brief methods overview

Under the standard topic model, we have

$$c_{ng} \sim Poi(c_{n+} \sum_{k=1}^{K} \omega_{nk} \theta_{kg})$$

Let us define

$$\lambda_{ng} = c_{n+} \sum_{k=1}^{K} \omega_{nk} \theta_{kg}$$

Under the normal model, if λ_{ng} is large, we can assume

$$c_{ng} \sim N(\lambda_{ng}, \lambda_{ng})$$

which is equivalent to saying

$$c_{ng} = \lambda_{ng} + e_{ng}$$
 $e_{ng} \sim N(0, \lambda_{ng})$

For applying *flash*, we first estimate the λ in the variance using topic model estimate (we call this λ to be λ^*).

to guard against the odd possibility that for some n and g, this estimate λ^* could be 0, we replace such cases as of now with a small value 0.0001.

In this example model we have considered $c_{n+} = 1000$.

```
lambda_star <- 1000 * (omega %*% t(theta))</pre>
lambda_star[lambda_star == 0] <- 1e-04</pre>
dim(lambda_star)
```

[1] 600 100

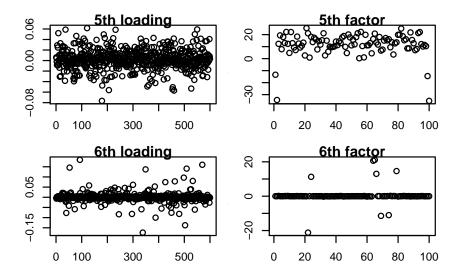
FLASH model fitting

flash then fits the model

$$c_{ng} = \sum_{k=1}^{K} l_{nk} f_{kg} + e_{ng}$$
 $e_{ng} \sim N(0, \lambda_{ng}^{\star})$

```
g1 = suppressMessages(flash(counts, sigmae2_true = lambda_star))
f = g1\$f
l = g1$l
res = counts - 1 %*% t(f)
# g_new = flash(res,nonnegative =
# TRUE, sigmae2_true = lambda)
g2 = suppressMessages(flash(res, sigmae2_true = lambda_star))
l = g2$l
f = g2\$f
res = res - l %*% t(f)
g3 = suppressMessages(flash(res, sigmae2_true = lambda_star))
l = g3$l
f = q3$f
res = res - l %*% t(f)
g4 = suppressMessages(flash(res, sigmae2_true = lambda_star))
l = g4$l
f = q4$f
res = res - l %*% t(f)
g5 = suppressMessages(flash(res, sigmae2_true = lambda_star))
l = g5$l
f = g5$f
res = res - l %*% t(f)
g6 = suppressMessages(flash(res, sigmae2_true = lambda))
l = g6$l
f = q6$f
par(mfrow = c(2, 2))
par(cex = 0.6)
par(mar = c(3, 3, 0.8, 0.8), oma = c(1, 1, 1,
    1))
plot(q1$l, main = "1st loading")
plot(g1$f, main = "1st factor")
```

```
plot(g2$l, main = "2nd loading")
plot(g2$f, main = "2nd factor")
                                                    1st factor
              1st loading
  -0.038
                                       -1000
  -0.046
                                       -2500
       0
          100
                   300
                            500
                                            0
                                                 20
                                                      40
                                                           60
                                                                80
                                                                     100
                                       2000
              2nd loading
                                                    2nd factor
  0.00
  0.00
                                       0
  -0.06
                                       -2000
       0
          100
                   300
                            500
                                            0
                                                 20
                                                      40
                                                           60
                                                                80
                                                                     100
plot(g3$l, main = "3rd loading")
plot(g3$f, main = "3rd factor")
plot(g4$1, main = "4th loading")
plot(g4$f, main = "4th factor")
                                                    3rd factor
              3rd loading
                                       100
  -0.10 0.05
                                       0
                                       -100
       0
          100
                   300
                            500
                                            0
                                                 20
                                                      40
                                                           60
                                                                80
                                                                     100
              4th loading
                                                    4th factor
                                       150
  0.00
                                       20
                                       -50
  -0.15
       0
          100
                   300
                            500
                                                      40
                                                                     100
                                            0
                                                 20
                                                           60
                                                                80
plot(g5$l, main = "5th loading")
plot(g5$f, main = "5th factor")
plot(g6$l, main = "6th loading")
plot(g6$f, main = "6th factor")
```

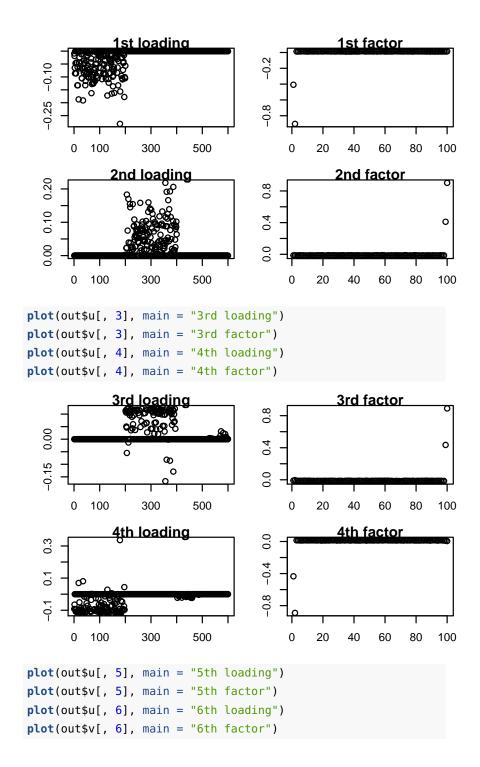


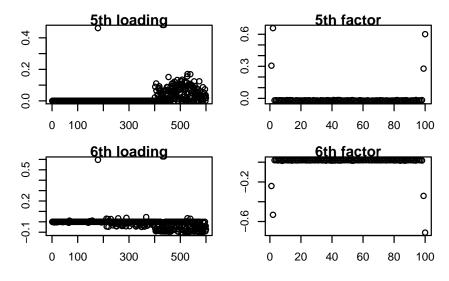
PMA model fitting

We compare the output of *flash* with that of the PMD() function of the package PMA. We use their default settings for shrinkage.

```
out <- PMD(counts, K = 6)
## 1234567
## 123456
## 12345678
## 12345678
## 123456789101112131415
## 123456789101112131415161718
par(mfrow = c(2, 2))
par(cex = 0.6)
par(mar = c(3, 3, 0.8, 0.8), oma = c(1, 1, 1,
    1))
plot(out$u[, 1], main = "1st loading")
plot(out$v[, 1], main = "1st factor")
plot(out$u[, 2], main = "2nd loading")
```

plot(out\$v[, 2], main = "2nd factor")





sessionInfo()

```
## R version 3.2.4 (2016-03-10)
## Platform: x86_64-apple-darwin13.4.0 (64-bit)
## Running under: OS X 10.10.5 (Yosemite)
##
## locale:
## [1] en_US.UTF-8/en_US.UTF-8/en_US.UTF-8/C/en_US.UTF-8/en_US.UTF-8
##
## attached base packages:
## [1] stats
                 graphics
                           grDevices utils
## [5] datasets methods
                           base
##
## other attached packages:
## [1] Rcpp_0.12.3
                     PMA_1.0.9
                                    impute_{-1.44.0}
## [4] plyr_1.8.3
                     irlba_2.0.0
                                   Matrix_1.2-4
## [7] ashr_1.0.8
##
## loaded via a namespace (and not attached):
   [1] knitr_1.12.3
                          magrittr_1.5
   [3] MASS_7.3-43
                          doParallel_1.0.10
##
   [5] pscl_1.4.9
                          SQUAREM_2014.8-1
##
  [7] lattice_0.20-33
                          foreach_1.4.3
##
                          tools_3.2.4
   [9] stringr_1.0.0
## [11] parallel_3.2.4
                          grid_3.2.4
                          htmltools_0.3
## [13] maptpx_1.9-2
## [15] iterators_1.0.8
                          assertthat_0.1
## [17] yaml_2.1.13
                          digest_0.6.9
## [19] formatR_1.2.1
                          codetools_0.2-14
## [21] slam_0.1-32
                          evaluate_0.8
```

[25] truncnorm_1.0-7 tufte_0.2