

# *flashtpx: Simulation Run 2*

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## *Overview*

This is the second 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 with 2 clusters with cluster membership in cluster 1 varying from 0.8 to 0.2 linearly.

```
n.out <- 600
omega_sim <- 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 2
```

```
K <- dim(omega_sim)[2]

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)
```



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],
  function(x) rmultinom(1, 1000, prob = omega_sim[x,
  ] %*% freq))))
dim(counts)
```

```
## [1] 600 100
```

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

[illegible]

### 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  $\lambda_{ng}$  is large, we can assume

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

which is equivalent to saying

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

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

to guard against the odd possibility that for some  $n$  and  $g$ , this estimate  $\lambda^*$  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))
lambda_star[lambda_star == 0] <- 1e-04
dim(lambda_star)
```

```
## [1] 600 100
```

### FLASH model fitting

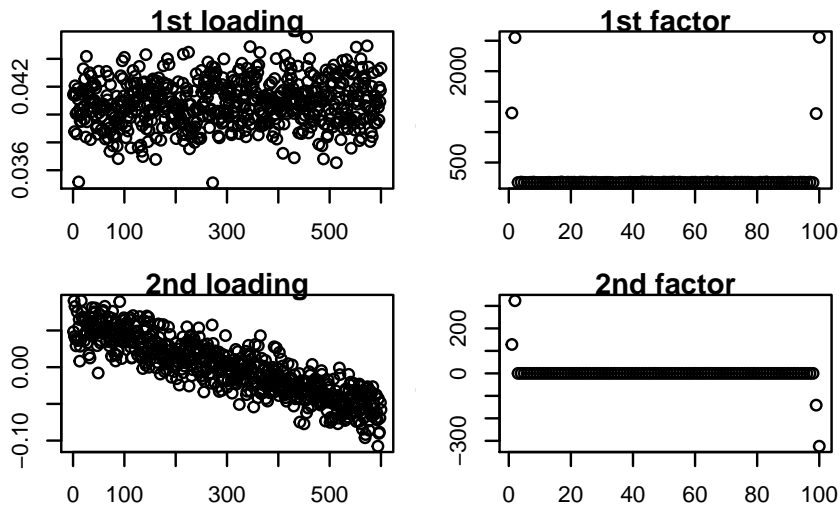
*flash* then fits the model

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

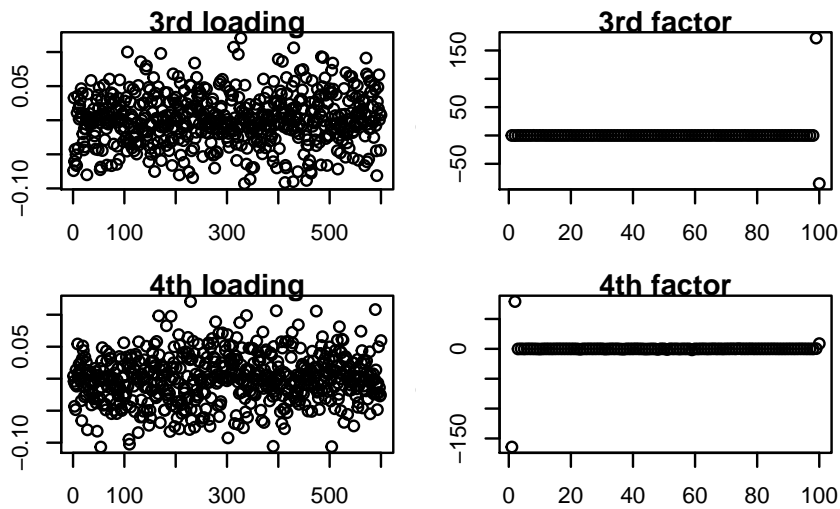
```

g1 = suppressMessages(flash(counts, sigmae2_true = lambda_star))
f = g1$f
l = g1$l
res = counts - l %*% 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$l
f = g3$f
res = res - l %*% t(f)
g4 = suppressMessages(flash(res, sigmae2_true = lambda_star))
l = g4$l
f = g4$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_star))
l = g6$l
f = g6$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(g1$l, main = "1st loading")
plot(g1$f, main = "1st factor")
plot(g2$l, main = "2nd loading")
plot(g2$f, main = "2nd factor")

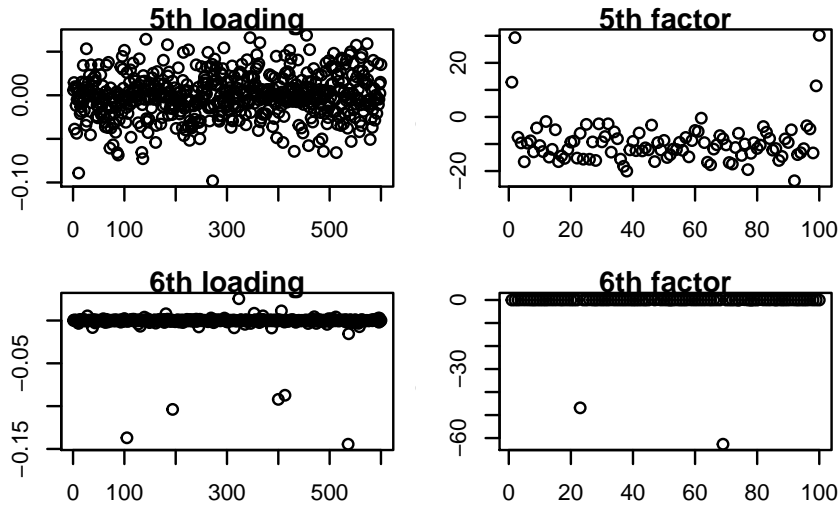
```



```
plot(g3$l, main = "3rd loading")
plot(g3$f, main = "3rd factor")
plot(g4$l, main = "4th loading")
plot(g4$f, main = "4th factor")
```



```
plot(g5$l, main = "5th loading")
plot(g5$f, main = "5th factor")
plot(g6$l, main = "6th loading")
plot(g6$f, main = "6th factor")
```



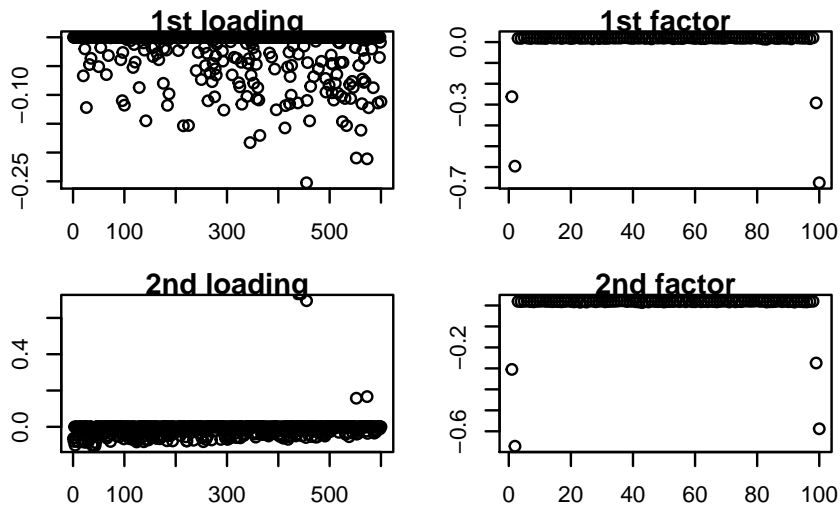
### *PMA model fitting*

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

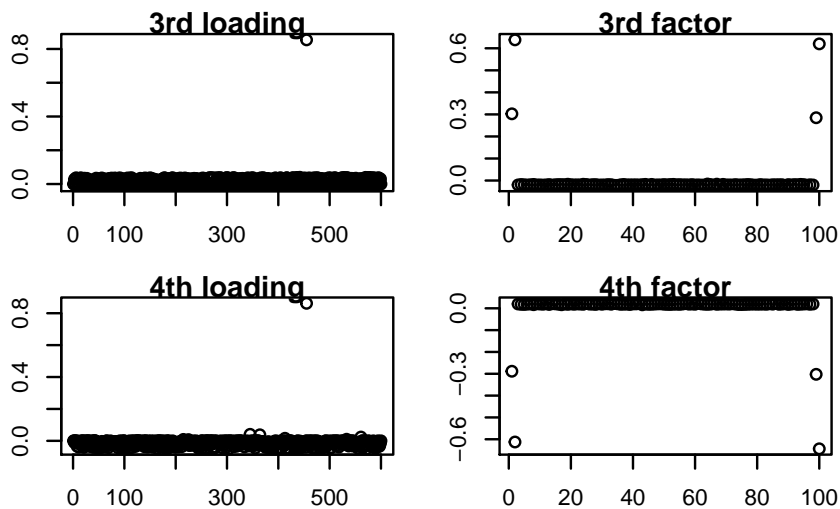
```
out <- PMD(counts, K = 6)
```

```
## 12345678910111213141516171819
## 1234567891011121314151617181920
## 12345678
## 123456789
## 12345678910111213
## 1234567891011121314151617181920
```

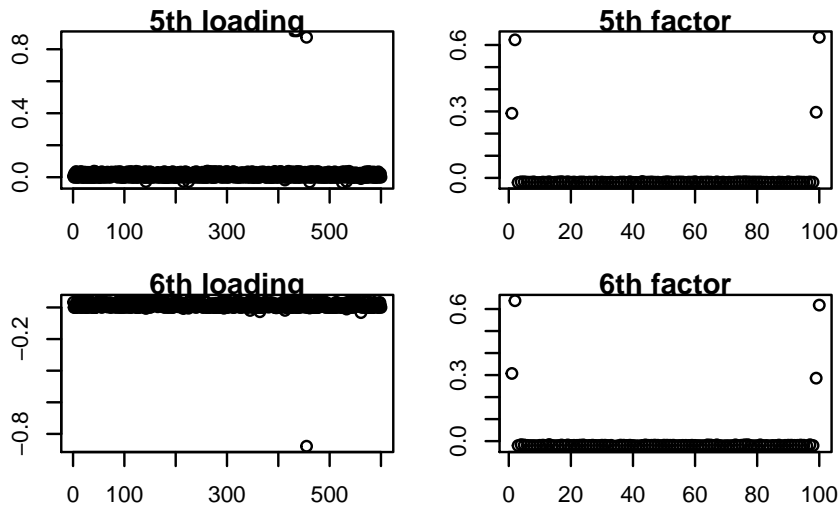
```
par(mfrow = c(2, 2))
par(cex = 0.6)
par(mar = c(3, 3, 0.8, 0.8), oma = c(1, 1, 1,
  1))
plot(out$u[, 1], main = "1st loading")
plot(out$v[, 1], main = "1st factor")
plot(out$u[, 2], main = "2nd loading")
plot(out$v[, 2], main = "2nd factor")
```



```
plot(out$u[, 3], main = "3rd loading")
plot(out$v[, 3], main = "3rd factor")
plot(out$u[, 4], main = "4th loading")
plot(out$v[, 4], main = "4th factor")
```



```
plot(out$u[, 5], main = "5th loading")
plot(out$v[, 5], main = "5th factor")
plot(out$u[, 6], main = "6th loading")
plot(out$v[, 6], main = "6th 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_0.12.4    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
## [9] stringr_1.0.0   tools_3.2.4
## [11] parallel_3.2.4  grid_3.2.4
## [13] maptpx_1.9-2    htmltools_0.3
## [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
```



```
## [23] rmarkdown_0.9.2  stringi_1.0-1  
## [25] truncnorm_1.0-7  tufte_0.2
```