

# Double shrinkage empirical Bayesian estimation for unknown and unequal variances

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In this paper, we construct a point estimator when assuming unequal and unknown variances by using the *empirical* Bayes approach in the classical normal mean problem. The proposed estimator shrinks both means and variances, and is thus called the double shrinkage estimator. Extensive numerical studies indicate that the double shrinkage estimator has lower Bayes risk than the estimator which shrinks the means alone, and the naive estimator which has no shrinkage at all. We further use a spike-in data set to assess different estimating procedures. It turns out that our proposed estimator performs the best and is thus strongly recommended for applications.

AMS 2000 SUBJECT CLASSIFICATIONS: Primary 60K35, 60K35; secondary 60K35.

KEYWORDS AND PHRASES: James–Stein estimator, Log-normal model, Loss function.

## 1. INTRODUCTION

The shrinkage estimator has a long history dating back to the 1950s. Assume that  $X_i \stackrel{\text{iid}}{\sim} N(\theta_i, \sigma_i^2)$  ( $i = 1, 2, \dots, p$ ). When  $p \geq 3$  and variances  $\sigma_i^2$ 's are known and all equal to  $\sigma^2$ , James and Stein (1961) proposed an estimator

$$(1) \quad \hat{\theta}_{JS} = \left(1 - \frac{(p-2)\sigma^2}{\sum_i X_i^2}\right) X$$

which dominates the estimator  $\hat{\theta} = X$  when using squared error loss

$$(2) \quad L(\hat{\theta}, \theta) = \frac{1}{p} \sum_i (\hat{\theta}_i - \theta_i)^2.$$

Further, it is easily seen that the estimator (1) can be dominated by the positive part James–Stein estimator defined as

$$(3) \quad \hat{\theta}_{JS+} = \left(1 - \frac{(p-2)\sigma^2}{\sum_i X_i^2}\right)_+ X = \delta_{JS+} X.$$

When simultaneously estimating all the parameters  $\theta_i$ 's, it is beneficial to use  $X_i$  and the seemingly unrelated observations  $X_{(-i)}$ , where the  $X_{(-i)}$  consists of all the observations  $X$  but  $X_i$ , when estimating each individual  $\theta_i$ . This is known

as borrowing strength effect. Although, the shrinkage scaling  $\delta_{JS+} = (1 - \frac{(p-2)\sigma^2}{\sum_i X_i^2})_+$  falls between zero and one, leading to biased estimators  $\hat{\theta}_{JS+}$  for all the parameters  $\theta_i$ 's, such scaling reduces the variances of the estimators resulting in a lower risk compared with the naive estimator  $X_i$ 's.

The positive part James–Stein's estimator is purely frequentist based. Lindley (1962) derived an estimator based on the *empirical* Bayesian approach described as following. Given a prior distribution of  $\theta_i$  as  $N(\mu, \tau^2)$ , the posterior expectation of  $\theta_i$  given  $X_i$  is  $MX_i + (1-M)\mu$  where  $M = \frac{\tau^2}{\tau^2 + \sigma^2}$  which minimizes the Bayes risk. Since  $\mu$  and  $\tau^2$  are usually unknown in practice, one estimates  $M$  and  $\mu$  from the data by using, for example, the method of moments. This resulted in the Lindley–James–Stein's estimator, abbreviated below as LJS in short,

$$\hat{\theta}_{LJS,i} = \hat{M}X_i + (1 - \hat{M})\bar{X},$$

where  $\hat{M}$  is a constant  $(1 - \frac{(p-3)\sigma^2}{\sum_i (X_i - \bar{X})^2})$  for all  $\theta_i$ 's. Since  $M$  is never negative, it is natural to further take the positive part of  $\hat{M}$  and obtain the positive part LJS estimator. This estimator is a shrinking-mean estimator. It pulls the observation  $X_i$  towards the arithmetic mean  $\bar{X}$ . It can also be shown that the positive part LJS has a smaller frequentist risk than that of LJS which in turn has smaller risk than  $X$  for every  $\theta$ 's.

The happy marriage of the James–Stein's idea and *empirical* Bayes approach brings a revolution in mathematical statistics. Statisticians use these ideas to produce different testing procedures, confidence intervals and others. To name a few, when assuming  $\vec{X} \sim N(\vec{\theta}, \sigma^2 I)$ , Casella and Hwang (1983) constructed a confidence set which dominates the naive confidence sphere centered at  $\vec{X}$  with radius  $c\sigma$ , where  $P(\chi_p^2 < c^2) = 1 - \alpha$ . Morris (1983a), Morris (1983b), and He (1992) constructed different *empirical* Bayesian confidence intervals for  $\theta_i$ 's. Qiu and Hwang (2007) constructed confidence intervals for selected  $\theta_i$ 's when assuming a mixed prior. See Casella and Hwang (2010) for a review.

However, all the literature listed above either assume a known variance  $\sigma^2$  or simply replace them by  $S_i^2$  when  $\sigma_i^2$ 's are unknown and unequal (See also Efron and Morris (1973); Morris (1983b)). This straightforward substitution results in a point estimator which only has the shrinking-mean effect for the heteroscedasticity case where the  $\sigma_i^2$ 's

are unequal and unknown. Fourdrinier et al. (2003) considered this type of estimators and proved that the frequentist risk under some weighted squared error of the new estimator is small. As we know, shrinking the means brings us much benefit, such as low risks in point estimators, shortness in intervals and powerfulness in testing procedures. What about shrinking the variances? Unlike shrinking the means, it is not until recently that researchers realize the advantage of the variance shrinkage and know how to shrink the variances in the context of estimating means.

Nowadays, in microarray experiments when the dimension  $p$  is very large, typically 10 thousands or more, some of the observations  $S_i^2$ 's can be either extremely large or small. In the spike-in data set we analyze in Section (5), the smallest value of  $S_i^2$ 's is  $6.0611 \times 10^{-5}$  while the largest one is 5.4160. Consequently, the testing procedure could either be of little power or detect much false significance. It seems advantageous to shrink the variances toward the common mean. This would enlarge the small variances and reduce the large variances.

Imposing an *inverse Gamma* prior of  $\sigma_i^2$  with hyper parameter  $a$  and  $b$  (see Berger (1985)) along with the assumption that  $\frac{S_i^2}{\sigma_i^2} \sim \frac{\chi_d^2}{d}$ , Smyth (2004) developed a better testing procedure by using the *empirical Bayes* approach. In Smyth's procedure, the variance  $\sigma_i^2$  is estimated by

$$(4) \quad \hat{\sigma}_i^2 = \frac{\frac{1}{b} + \frac{dS_i^2}{2}}{\frac{d}{2} + \hat{a} - 1}$$

where  $\hat{a}$  and  $\hat{b}$  are estimated by a numerical algorithm. This estimator truncates the small value of  $S_i^2$  to be at least  $\frac{1/\hat{b}}{d/2 + \hat{a} - 1}$ . When  $S_i^2 \gg \frac{1}{b}$ ,  $\hat{\sigma}_i^2$  is at most  $\frac{d/2}{d/2 + \hat{a} - 1} S_i^2$  which is smaller than  $S_i^2$  given  $\hat{a} > 1$ . Consequently, Smyth's variance estimator shrinks the variances. However, there are no explicit formulas of  $\hat{a}$  and  $\hat{b}$ , and it is hard to evaluate the property of the estimator analytically.

In 2005, Cui et al. (2005) proposed the exponential LJS estimator for the variance component  $\sigma_i^2$ 's with an explicit form which shrinks the observation towards their geometric mean as explained in Section 4. They further argued that the testing procedure  $F_S$  based on this variance shrinkage estimator enjoys high power. The subscript  $S$  here means that the procedure has only one shrinkage factor – shrinking the variances.

In addition to modeling the true parameter  $\theta_i$ 's, Hwang and Liu (2010) further put a log-normal prior for the variances  $\sigma_i^2$ 's. After approximating the  $\frac{\chi_d^2}{d}$  by another log-normal random variable, they proposed the so-called *Log-Normal* model. Using the *empirical Bayes* approach, they derived another testing procedure  $F_{SS}$  where the subscript  $SS$  means that this procedure has double-shrinkage factor—shrinking both the means and variances. They have demonstrated that the average power of  $F_{SS}$  is higher than that of all the other tests, such as  $F_S$ , the shrinking variance alone

test, and the  $T$ -test without any shrinkage. They have further concluded that it is better than the moderated  $T$ -test based on the variance shrinkage estimator (4) as in Smyth (2004).

Based on the same model, Hwang, Qiu and Zhao (2009) studied the *empirical Bayes* confidence interval with the double shrinkage effect. It turns out that this new construction dominates the naive  $t$  interval in terms of a sharper average length when guaranteeing the *empirical Bayes* coverage probability. They further argued that the confidence interval with double shrinkage is better than both the shrinking-mean-alone and the shrinking-variance-alone interval, which are better than intervals with no shrinkage.

It is interesting that no study aims at constructing estimators that shrink both the means and variances as far as the author is aware. This is what we do in this paper. We construct estimators for  $\theta$ 's which shrinks both the means and variances. The article is organized as following. In Section 2, we introduce the *general Log-Normal* model and derive a point estimator when assuming known hyperparameters. In Section 3, we estimate the hyper parameters from the data and derive the *empirical Bayes* estimator. We study the Bayes risk of the new estimator for a class of priors based on the loss function (2) by using extensive simulation studies and a real data analysis in Sections 4 and 5. We conclude in Section 6 that the point estimator with the double shrinkage is better than the estimator with one shrinkage, which is better than the estimator without any shrinkage.

## 2. ESTIMATOR WITH KNOWN HYPER PARAMETER

In this section, we define the canonical model over which we shall construct double shrinkage estimators. Firstly, assume that each observation  $X_i$  ( $i = 1, \dots, p$ ) follows a normal distribution with mean  $\theta_i$  and unknown variance  $\sigma_i^2$  which differ across all the observations. The heteroscedasticity of  $\sigma_i^2$  occurs often in application but causes lots of difficulties. We assume that there exists another statistic  $S_i^2$  containing the information of the variance  $\sigma_i^2$  which is independent of  $X_i$ . In general, it is assumed that  $S_i^2 | \sigma_i^2 \stackrel{\text{iid}}{\sim} \sigma_i^2 \frac{\chi_{d_i}^2}{d_i}$  where  $d_i$  represents the degrees of freedom corresponding to the  $i$ -th observation.

In modern application such as microarray technology, the dimension  $p$  is large, typically varying from several thousands to 50 thousands. Therefore, it is practical to put a prior distribution over  $\theta_i$ . When assuming that all the  $\sigma_i^2$ 's are equal and known as  $\sigma^2$ , Lindley (1962) assumes a normal prior  $N(\mu, \tau^2)$  for  $\theta_i$  and derived the well known Lindley James–Stein estimator of  $\theta_i$  as

$$(5) \quad \hat{\theta}_i = \bar{X} + \left(1 - \frac{(p-3)\sigma^2}{\sum_i (X_i - \bar{X})^2}\right) (X_i - \bar{X}).$$

Similarly, in our model, we assume the same prior  $N(\mu, \tau^2)$  for the true parameter  $\theta_i$ .

There are many variances  $\sigma_i^2$ 's, and it seems reasonable to assume a prior for the variances  $\sigma_i^2$ 's as well. It is convenient to put an inverse gamma prior with the shape parameter  $a$  and scale parameter  $b$  (see Berger (1985)) for  $\sigma_i^2$  because it is conjugate to the  $\chi^2$  random variable. Smyth (2004) took this approach and derived an *empirical* Bayes testing procedure. However, a disadvantage of such an approach is that there is no explicit formula for the estimator of the hyper parameters. Smyth (2004) introduced a numerical algorithm to estimate these two parameters.

In our model, we first approximate  $\log \frac{\chi_{d_i}^2}{d_i}$  by  $N(m_i, \sigma_{ch,i}^2)$  where

$$m_i = E \log \frac{\chi_{d_i}^2}{d_i} = \psi\left(\frac{d_i}{2}\right) - \log \frac{d_i}{2},$$

$$\sigma_{ch,i}^2 = Var\left(\log \frac{\chi_{d_i}^2}{d_i}\right) = \psi'\left(\frac{d_i}{2}\right),$$

where  $\psi(x) = \frac{d}{dx} \log \Gamma(x)$ , known as the *digamma* function. The two constants  $m_i$  and  $\sigma_{ch,i}^2$  depend solely on the degrees of freedom  $d_i$  and can be evaluated easily. Consequently, we approximate the logarithm of  $\frac{\chi_{d_i}^2}{d_i}$  by a normal random variable with the same first and second moments. As a result,

$$\log S_i^2 | \log \sigma_i^2 \stackrel{\text{iid}}{\sim} N(m_i + \log \sigma_i^2, \sigma_{ch,i}^2).$$

Furthermore, we assume that  $\log \sigma_i^2$  is a normal random variable with a hyper mean  $\mu_v$  and a variance  $\tau_v^2$ . To distinguish with the prior distribution of the mean  $\theta_i$ , we use the sub-index  $v$ , since they are the hyper parameters corresponding to the variances.

In summary, the canonical model is

$$(6) \quad \begin{cases} X_i | \theta_i, \sigma_i^2 \stackrel{\text{iid}}{\sim} N(\theta_i, \sigma_i^2); \\ \theta_i \stackrel{\text{iid}}{\sim} N(\mu, \tau^2); \\ \log S_i^2 | \log \sigma_i^2 \stackrel{\text{iid}}{\sim} N(m_i + \log \sigma_i^2, \sigma_{ch,i}^2); \\ \log \sigma_i^2 \stackrel{\text{iid}}{\sim} N(\mu_v, \tau_v^2), \end{cases}$$

where  $m_i = \psi\left(\frac{d_i}{2}\right) - \log \frac{d_i}{2}$ ,  $\sigma_{ch,i}^2 = \psi'\left(\frac{d_i}{2}\right)$ .

This model is called *Log-normal* model in Hwang and Liu (2010) and Hwang, Qiu and Zhao (2009) where they assume that the degrees of freedom  $d_i$  across all observations are the same. Hwang and Liu (2010) constructed a powerful testing procedure while Hwang, Qiu and Zhao (2009) constructed sharp *empirical* Bayesian confidence intervals based on the same model setting. In the data analysis part of Hwang, Qiu and Zhao (2009) when the degrees of freedom of each gene are either 2 or 3, they took a conservative approach and simply set all degrees of freedom to be 2. As illustrated in Section 4, if taking the same conservative approach when the degrees of freedom  $d_i$ 's are different, the corresponding estimator has a slightly larger risk when compared with a new point estimation procedure based on this Log-Normal model, where  $d_i$  are not necessarily identical.

Having the model, we first derive the point estimator  $\hat{\theta}$  when assuming that all the hyper-parameters  $\mu, \tau^2, \mu_v, \tau_v^2$  are known. Since  $X_i | \theta_i \sim N(\theta_i, \sigma_i^2)$  and  $\theta_i \sim N(\mu, \tau^2)$ , we know that  $\theta_i | X_i, \sigma_i^2 \sim N(M_i X_i + (1 - M_i)\mu, M_i \sigma_i^2)$  where  $M_i = \frac{\tau^2}{\tau^2 + \sigma_i^2}$ . For the known variance  $\sigma_i^2$  case, the natural estimator of  $\theta_i$  is  $M_i X_i + (1 - M_i)\mu$ , which is the posterior expectation of  $\theta_i$  given  $X_i$  and  $\sigma_i^2$ . This estimator shrinks the observation  $X_i$  towards the hyper mean  $\mu$ . The shrinkage scaling  $M_i$  equals  $\frac{\tau^2}{\tau^2 + \sigma_i^2}$ , which depends on the variance  $\sigma_i^2$  of the  $i$ -th observation.

However, since  $\sigma_i^2$  is unknown, we need to substitute it by a variance estimator  $\hat{\sigma}_i^2$  depending on the observation  $S_i^2$  and hyper parameter  $\mu_v$  and  $\tau_v^2$ . One typical approach is to replace  $\sigma_i^2$  by  $S_i^2$ , and estimate  $\theta_i$  by

$$\hat{\theta}_i = \hat{M}_i X_i + (1 - \hat{M}_i)\mu,$$

where  $\hat{M}_i = \frac{\tau_v^2}{\tau_v^2 + S_i^2}$ . This estimator shrinks the observation  $X_i$  towards the common mean  $\mu$ . However, there is no variance shrinkage.

Recall in model (6) one knows that,

$$\log S_i^2 | \log \sigma_i^2 \sim N(m_i + \log \sigma_i^2, \sigma_{ch,i}^2),$$

and

$$\log \sigma_i^2 \sim N(\mu_v, \tau_v^2).$$

A classical calculation indicates that

$$\log \sigma_i^2 | \log S_i^2 \sim N(M_{v,i}(\log S_i^2 - m_i) + (1 - M_{v,i})\mu_v, M_{v,i}\sigma_{ch,i}^2)$$

where  $M_{v,i} = \frac{\tau_v^2}{\tau_v^2 + \sigma_{ch,i}^2}$ . There exists two natural estimators of the  $\sigma_i^2$  based on the previous posterior density as

$$(7) \quad \begin{aligned} \hat{\sigma}_{i,1}^2 &= \exp(E \log \sigma_i^2 | S_i^2) \\ &= \exp(M_{v,i}(\log S_i^2 - m) + (1 - M_{v,i})\mu_v), \end{aligned}$$

and

$$(8) \quad \hat{\sigma}_{i,2}^2 = E(\sigma_i^2 | S_i^2) = \hat{\sigma}_{i,1}^2 \exp\left(\frac{M_{v,i}\sigma_{ch,i}^2}{2}\right).$$

The estimator  $\hat{\sigma}_{i,2}^2$  is based on the exact posterior expectation. When constructing a confidence interval for  $\theta_i$ , Hwang, Qiu and Zhao (2009) prefers the estimator  $\hat{\sigma}_{i,1}^2$  because it produces a shorter, in other words, more efficient interval than the other one when both guaranteeing the *empirical* Bayesian coverage probability. Here we will use the estimator  $\hat{\sigma}_{i,1}^2$ , written as  $\hat{\sigma}_i^2$ , to construct the estimator of  $\theta_i$  later in this paper. Practically speaking, there is little difference for these two approaches in estimating  $\theta_i$ 's.

The exact posterior distribution of  $\sigma_i^2$  given the observation  $(X_i, S_i^2)$  also depends on  $X_i$  and has no explicit form. We approximate this posterior by assuming that it depends

solely on  $S_i^2$ . This approximation is practically and intuitively reasonable.

Having the variance shrinkage estimator, we now turn to the estimation of  $\theta_i$ . Recall that

$$E(\theta_i|X_i, \sigma_i^2) = M_i X_i + (1 - M_i)\mu.$$

Then we can estimate  $\theta_i$  by

$$(9) \quad \hat{\theta}_i = \hat{M}_i X_i + (1 - \hat{M}_i)\mu,$$

where  $\hat{M}_i = \frac{\tau_v^2}{\tau_v^2 + \hat{\sigma}_i^2}$ .

The above estimator is not the exact Bayes estimator  $E(\theta_i|X_i, S_i^2)$  which minimizes the Bayes risk when using the loss function (2). However, (9) has the advantage of having instantaneous computation. It would be very interesting to derive some analytic results regarding the relations between (9) and the exact Bayes estimator.

### 3. ESTIMATING THE HYPER-PARAMETER

In Section 2, we have proposed the point estimator of  $\theta$  when assuming known hyper parameters  $\mu, \tau^2, \mu_v$  and  $\tau_v^2$ , which, in practice, are unknown. To avoid any subjective choice, we incorporate the empirical Bayes approach by estimating the parameters through the data. The estimation resembles the calculation in Hwang, Qiu and Zhao (2009) where the method of moments is used.

Firstly, we estimate the hyper parameters  $\mu_v, \tau_v^2$  corresponding to the variances component. In model (6), it is assumed that

$$\log S_i^2 - m_i | \log \sigma_i^2 \sim N(\log \sigma_i^2, \sigma_{ch,i}^2),$$

and

$$\log \sigma_i^2 \sim N(\mu_v, \tau_v^2).$$

Consequently,  $E(\log S_i^2 - m_i) = \mu_v$ . We estimate  $\mu_v$  by

$$\hat{\mu}_v = \frac{1}{p} \sum_i (\log S_i^2 - m_i).$$

Further,  $E(\log S_i - m_i)^2 = \mu_v^2 + \tau_v^2 + \sigma_{ch,i}^2$ . We thus estimate  $\tau_v^2$  by

$$\hat{\tau}_v^2 = \left( \frac{1}{p} \left( \sum_i (\log S_i^2 - m_i)^2 - \sigma_{ch,i}^2 - \hat{\mu}_v^2 \right) \right)_+,$$

and

$$\hat{M}_{v,i} = \frac{\hat{\tau}_v^2}{\hat{\tau}_v^2 + \sigma_{ch,i}^2}.$$

Providing with the estimation of the hyper parameter corresponding to the variances and formula (7), we derive an empirical Bayes estimator of  $\sigma_i^2$  as

$$(10) \quad \hat{\sigma}_{EB,i}^2 = \exp(\hat{M}_{v,i}(\log S_i^2 - m_i) + (1 - \hat{M}_{v,i})\hat{\mu}_v).$$

When the degrees of freedom  $d_i = d$  ( $i = 1, \dots, p$ ), the estimator (10) can be written as

$$\hat{\sigma}_{EB,i}^2 = \exp(\hat{M}_v(\log S_i^2 - m) + (1 - \hat{M}_v)\hat{\mu}_v),$$

where

$$m = \psi(d/2) - \log(d/2), \sigma_{ch}^2 = \psi'(d/2)$$

and

$$\hat{M}_v = \left( 1 - \frac{(p-3)\sigma_{ch}^2}{\sum_i (\log S_i^2 - \log S^2)^2} \right)_+$$

which is the exponential Lindley–James–Stein’s estimator introduced in Cui et al. (2005).

Note that the empirical Bayes estimator (10) can be written as

$$\hat{\sigma}_{EB,i}^2 = \left( \left( \prod_i \frac{S_i^2}{e^{m_i}} \right)^{1/p} \right)^{1-\hat{M}_v} \left( \frac{S_i^2}{e^{m_i}} \right)^{\hat{M}_v}.$$

This indicates that  $\hat{\sigma}_{EB,i}$  shrinks the observation  $S_i^2/\exp(m_i)$  towards their geometric mean  $(\prod_i S_i^2/\exp(m_i))^{1/p}$ , resulting in a variance shrinkage estimator.

The next step is to estimate the hyper parameters  $\mu$  and  $\tau^2$  of the means  $\theta_i$ ’s. Since

$$X_i | \sigma_i^2 \sim N(\mu, \sigma_i^2 + \tau^2),$$

we estimate  $\mu$  by the weighted average as

$$\hat{\mu} = \frac{\sum X_i / \hat{\sigma}_{EB,i}^2}{\sum 1 / \hat{\sigma}_{EB,i}^2}.$$

Further, since  $E(X_i - \mu)^2 | \sigma_i^2 = \sigma_i^2 + \tau^2$ , Hwang, Qiu and Zhao (2009) estimated  $\tau^2$  as

$$\hat{\tau}^2 = \left( \frac{\sum (X_i - \hat{\mu})^2 - \hat{\sigma}_{EB,i}^2}{p} \right)_+.$$

However, the estimator  $\hat{\sigma}_{EB,i}^2$  is not an unbiased estimator of  $\sigma_i^2$ , resulting in an inconsistent estimator of  $\tau^2$  as  $p \rightarrow \infty$ . In order to remedy this, we estimate  $\tau^2$  by using

$$\hat{\tau}^2 = \left( \frac{\sum (X_i - \hat{\mu})^2 - S_i^2 \exp(-m_i - \sigma_{ch,i}^2/2)}{p} \right)_+,$$

due to the fact that

$$E S_i^2 | \log \sigma_i^2 = \sigma_i^2 \exp \left( m_i + \frac{\sigma_{ch,i}^2}{2} \right).$$

When assuming that  $\frac{S_i^2}{\sigma_i^2} \sim \chi_{d_i}^2$ , other than the log-normal assumption, we remove the term  $\exp(-m_i - \sigma_{ch,i}^2/2)$  when

estimating  $\tau^2$  in order to obtain a consistent estimator of  $\tau^2$ .

With the estimators of all the hyper parameters available, we propose the estimator for  $\theta_i$  as

$$(11) \quad \hat{\theta}_{SS,i} = \hat{M}_{EB,i} X_i + (1 - \hat{M}_{EB,i}) \hat{\mu},$$

where  $\hat{M}_{EB,i} = \frac{\hat{\tau}^2}{\hat{\tau}^2 + \sigma_{EB,i}^2}$ .

It is worthy noting that the estimator (11) is a shrinking-mean estimator for it shrinks the observation  $X_i$  towards the weighted average  $\hat{\mu}$ . Additionally, the estimator  $\sigma_{EB,i}^2$ , as defined in (10), is a variance shrinkage estimator as it shrinks the observation  $S_i^2 / \exp(\mu_i)$ 's towards their geometric mean. Therefore, we call the estimator (11) as double shrinkage estimator  $\hat{\theta}_{SS}$ . When estimating  $\theta_i$ , especially the hyperparameters  $\mu$ ,  $\mu_v$ ,  $\tau^2$ , and  $\tau_v^2$ , we borrow the strength from the seemingly unrelated observations  $X_j$ , and  $S_j^2$  where  $j \neq i$ .

#### 4. SIMULATION STUDY

In Section 3, we have proposed the double shrinkage estimator  $\hat{\theta}_{SS}$  of  $\theta$  which shrinks both the means and variances. Alternatively, if replacing the variance  $\sigma_i^2$  simply by  $S_i^2$  and through replicating the procedure above, one can propose an alternative estimator  $\hat{\theta}_{SM,i}$  as

$$(12) \quad \hat{\theta}_{SM,i} = \hat{M}'_{EB,i} X_i + (1 - \hat{M}'_{EB,i}) \hat{\mu}',$$

where  $\hat{M}'_{EB,i}$  and  $\hat{\mu}'$  are derived similarly as in Section 3 with  $\sigma_{EB,i}^2$  replaced by  $S_i^2$ . Such an estimator is called shrink-mean-alone estimator for it shrinks  $X_i$  towards the weighted average  $\hat{\mu}$  and has no variance shrinkage. Like  $\hat{\theta}_{SS,i}$ , this estimator also has the borrowing strength effect.

In addition, one can estimate  $\theta$  simply by  $\hat{\theta}_{NS} = X$  which neither shrinks nor borrows strength from other observations. In this section, we use simulation studies to calculate the Bayes risk under various parameter settings and model settings. The loss function is defined in (2).

In Figures 1 and 3, random numbers are generated according to the genuine *Log-Normal* model. We have simulated the Bayes risk of the estimators  $\hat{\theta}_{SS}$  as in (11),  $\hat{\theta}_{SM}$  as in (12), and  $\hat{\theta}_{NS} = X$  with the dimension  $p$  being 2,000. Their risks are represented by curves with markers as Diamonds, Circles, and Crossings respectively.

In Figure 1, the degrees of freedom  $d_i$ 's are randomly selected among 2, 3, 4 and 5. The hyper parameters  $\mu = \mu_v = 0$  and  $\tau_v^2$  varies among 0, 0.25, 0.5, and 1 from the top to the bottom. The Bayes risk is plotted against  $M = \frac{\tau^2}{\tau^2 + \exp(\mu_v + \sigma_v^2/2)}$ , varying from 0 to  $\infty$ . In other words, the hyper parameter  $\tau^2$  goes from 0 to  $\infty$ . In Figure 3, all the degrees of freedom  $d_i$ 's equal 2.

From these two figures, it is seen that  $\hat{\theta}_{SS}$  always dominates both  $\hat{\theta}_{SM}$  and  $\hat{\theta}_{NS}$  for different hyper parameter settings. Both the shrinkage estimators substantially improve

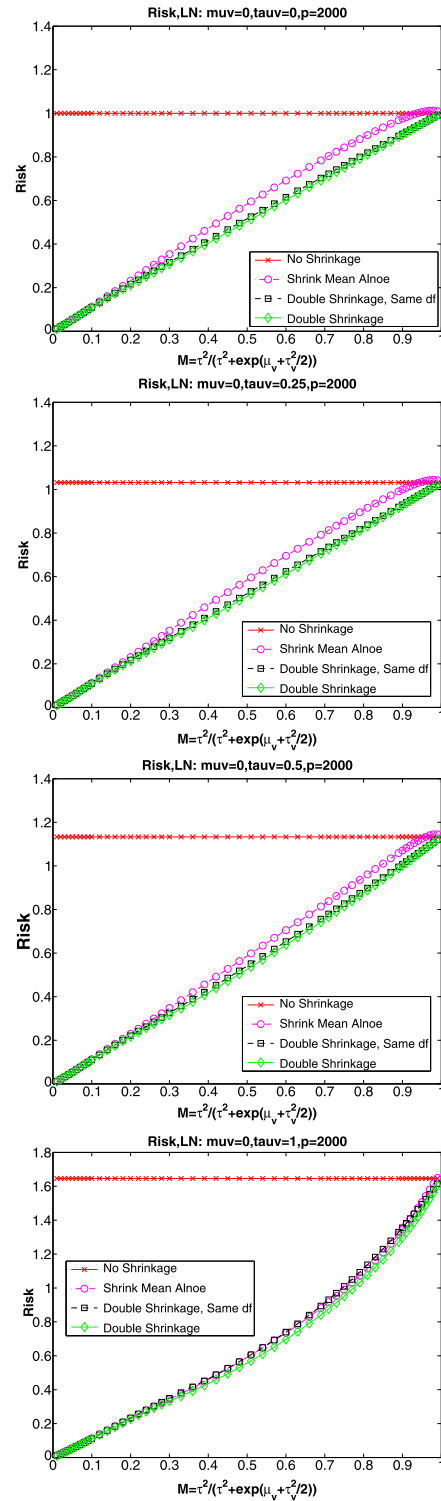


Figure 1. These figures are the Bayes risks of four point estimators with the dimension  $p = 2,000$ . The random numbers are generated according to the genuine *Log-Normal* model. The degrees of freedom are randomly chosen from 2 to 5. The hyper parameter setting are  $\mu = 0$ ,  $\mu_v = 0$ . The  $\tau_v^2$  varies from 0, 0.25, 0.5, to 1 from the top to the bottom. We plot the risk against  $M = \frac{\tau^2}{\tau^2 + \exp(\mu_v + \tau_v^2/2)}$  which goes from 0 to 1.



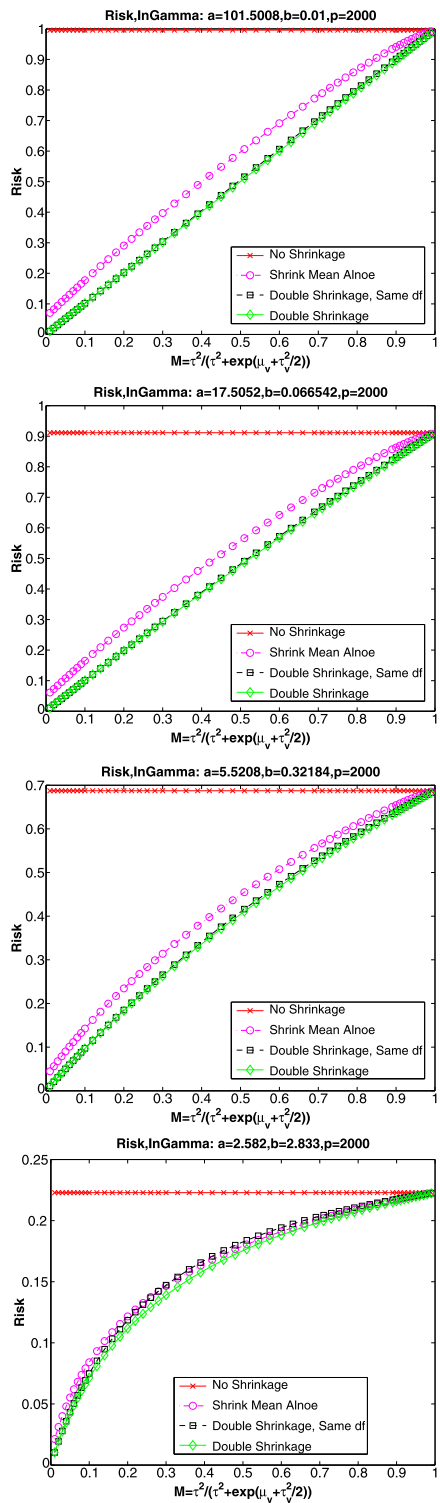


Figure 2. These figures are the Bayes risks of four point estimators with the dimension  $p = 2,000$ . The degrees of freedom are randomly chosen among 2, 3, 4, and 5. The random numbers are generated according to the inverse gamma model. The hyper parameters  $a$  and  $b$  are chosen according to (14). We plot the risk against  $M$  which goes from 0 to 1. The hyper parameter  $\mu$  is 0.

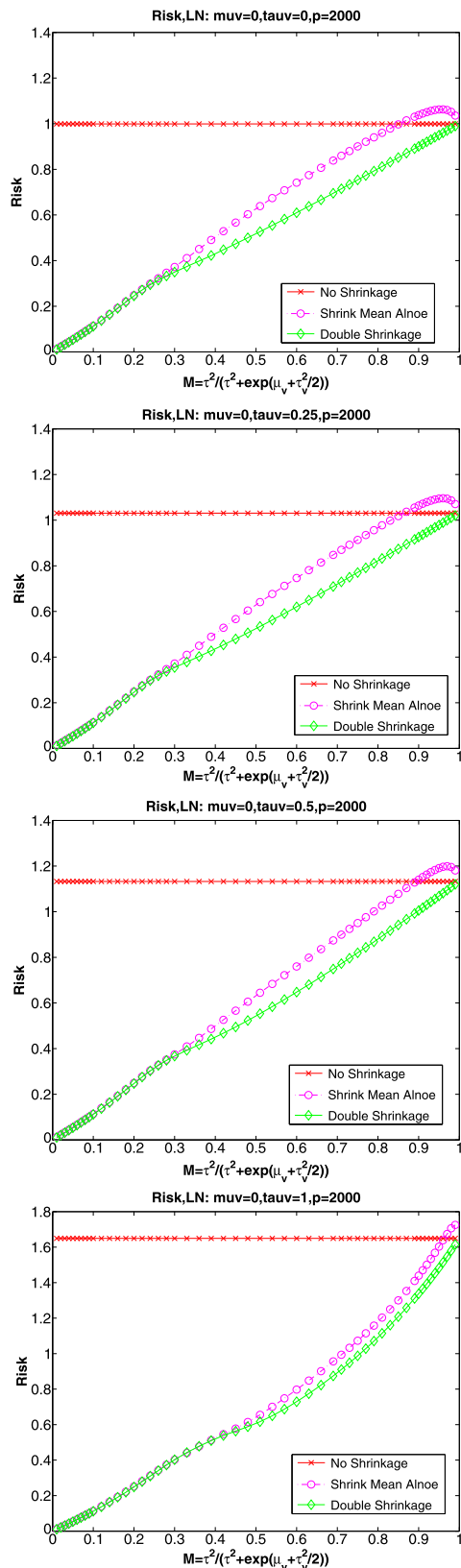


Figure 3. The parameter settings are the same as that in Figure 1. The only difference is that the degrees of freedom  $d_i$ 's are the same and equal to 2.

$\hat{\theta}_{NS}$  when  $\tau^2$  is close to 0. This indicates that shrinking-mean is important when all the true means are close. On the other hand, when  $\tau^2$  goes to infinity, the Bayes risk of double shrinkage estimator converges to the risk of  $\hat{\theta}_{NS}$  from below. Surprisingly, the Bayes risk of  $\hat{\theta}_{SM}$  exceeds the level of that of the no shrinkage estimator  $X$  for large  $\tau^2$  and small degrees of freedom. We further notice that  $\hat{\theta}_{SS}$  dominates  $\hat{\theta}_{SM}$  under every case. The improvement is significant especially for small  $\tau^2$  when the variances  $\sigma_i^2$ 's are close to each other. When  $\tau_v^2$  is large,  $\hat{\theta}_{SM}$  performs nearly the same as  $\hat{\theta}_{SS}$ .

In conclusion, the simulation results show that  $\hat{\theta}_{SS}$  dominates both  $\hat{\theta}_{SM}$  and  $\hat{\theta}_{NS}$  under the log-normal model.

In Figures 2 and 4, we have generated the random number according to the *inverse gamma* model with the last two equation of model (6) being replaced by

$$(13) \quad \begin{cases} S_i^2 | \sigma_i^2 \sim \sigma_i^2 \frac{\chi_{d_i}^2}{d_i}; \\ \sigma_i^2 \sim \text{Inverse Gamma}(a, b). \end{cases}$$

In other words,  $(\sigma_i^2)^{-1}$  has a *Gamma* distribution with parameters  $a$  and  $b$ . See Berger (1985).

In these simulations, the dimension  $p = 2,000$ . The degrees of freedom  $d_i$ 's are randomly chosen among 2, 3, 4, and 5 in Figure 2 and set to be 2 in Figure 4. The hyper parameters  $a$  and  $b$  are chosen such that

$$(14) \quad E\sigma_i^2 = E(\exp(N(\mu_v, \tau_v^2))), \text{Var}(\sigma_i^2) = \text{Var}(\exp(N(\mu_v, \tau_v^2)))$$

where  $\mu_v = 0$ ,  $\tau_v^2$  varies among 0, 0.25, 0.5, and 1 from the top to the bottom in each figure.

In all these studies, the Bayes risk of  $\hat{\theta}_{SS}$  is smaller than that of  $\hat{\theta}_{SM}$ , which is smaller than that of  $\hat{\theta}_{NS}$ . The improvement of  $\hat{\theta}_{SM}$  over  $\hat{\theta}_{NS}$  is very substantial for small  $\tau^2$ . When  $\tau^2 \rightarrow \infty$ , in other words,  $M \rightarrow 1$ , the Bayes risk of the shrinkage estimators converge to the risk of no shrinkage estimator from below. For small  $\tau_v^2$ , the double shrinkage estimator improves shrink-mean-alone estimator especially for small degrees of freedom. In Figure 5, we have simulated the Bayes risk of the estimators based on the *inverse gamma* model with  $\tau_v^2 = 0$  and equal degrees of freedom  $d$ , which varies among 2, 6, 10, and 20. The discrepancy between  $\hat{\theta}_{SS}$  and  $\hat{\theta}_{SM}$  gets smaller when the degrees of freedom increases. Nevertheless,  $\hat{\theta}_{SS}$  always dominates  $\hat{\theta}_{SM}$ .

In both Figures 1 and 2 when the degrees of freedom  $d_i$ 's are different across the observations, we have plotted the risk of the double shrinkage estimator when simply putting all the degrees of freedom to be the  $\min_{1 \leq i \leq p} d_i$ . This approach was taken by Hwang, Qiu and Zhao (2009) in constructing the confidence interval for each parameter  $\theta_i$ . The Bayes risk of this estimator is represented by the lines with Squares in these figures. It turns out that it is dominated by the new estimator  $\hat{\theta}_{SS}$ .

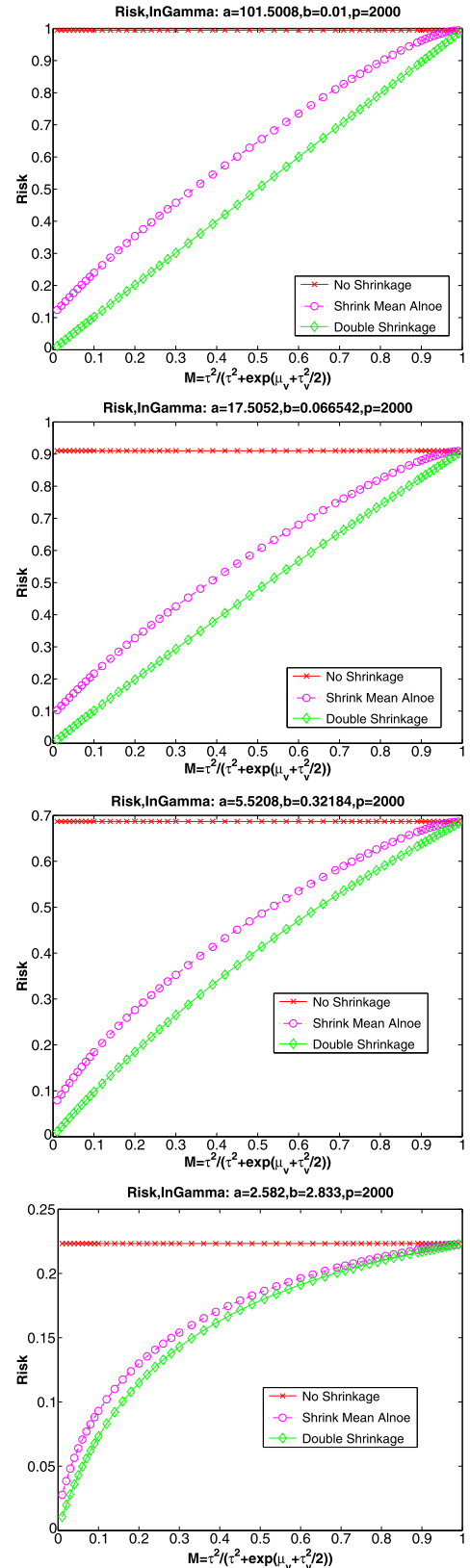


Figure 4. The parameter settings are the same as that in Figure 2. The only difference is that the degrees of freedom  $d_i$ 's are the same and equal to 2.

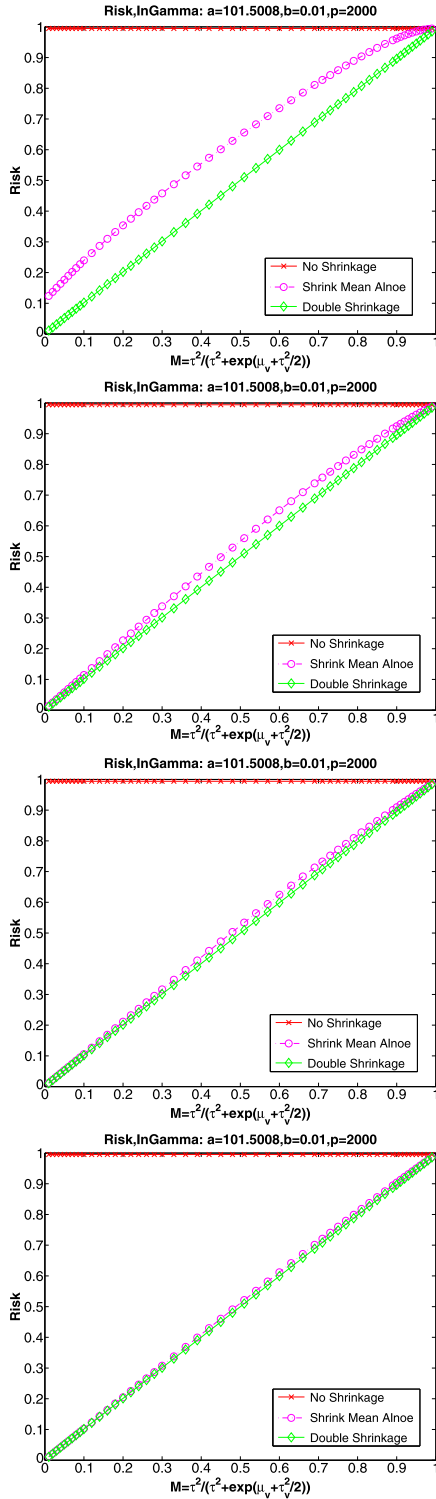


Figure 5. In this study, the data are generated according to the inverse Gamma model with  $p$  being 2,000. The hyper parameters  $\mu = \mu_v = 0, \tau_v = 0$ . The parameters  $a$  and  $b$  are chosen according to (14). In each graph, the degrees of freedom are the same and equal to 2, 5, 10, and 20 from the top to the bottom. The discrepancy between the risk of  $\hat{\theta}_{SS}$  and  $\hat{\theta}_{SM}$  increases when  $d$  decreases.

We have used sum of squared error loss (2) in the simulation. One of the referees pointed out that the same conclusion holds when we are only interested in estimating one parameter, saying  $\theta_1$ , with the one-dimensional squared error loss  $L(\theta_1, \hat{\theta}_1) = (\hat{\theta}_1 - \theta_1)^2$ . By claiming domination of the one-dimensional loss, the estimator we proposed is better than others under any weighted squared error loss.

All these simulation studies indicate that the double shrinkage estimator dominates the shrink-mean-alone estimator and no shrinkage estimator. In addition,  $\hat{\theta}_{SS}$  always performs the best for these two different model settings. This demonstrates that the new procedure is robust in some sense. Thus, the double shrinkage estimator  $\hat{\theta}_{SS}$  is strongly recommended.

## 5. REAL DATA ANALYSIS

We apply different estimators to an Affymetrix Control data set, the golden spike-in data set of Choe et al. (2005). All the parameters in this data set are pre-chosen and known. Therefore, it can be used to check different statistical procedures, such as the performance of confidence intervals in Hwang, Qiu and Zhao (2009) and point estimators as stated in this article.

In this section, we will calculate the risks of estimators  $\hat{\theta}_{SS}$ ,  $\hat{\theta}_{SM}$ , and  $\hat{\theta}_{NS}$ . We download the data from <http://www.elwood9.net/spike>. After taking the  $\log_2$  transformation, we fit the data to a one-way ANOVA model with the number of genes  $p$  being 14,010. There are 6 replicates for each gene, three from each of the control and treatment group. Let

$$X_i = \bar{Y}_{i1} - \bar{Y}_{i2}, S_i^2 = \sqrt{s_{1i}^2/3 + s_{2i}^2/3}.$$

The degrees of freedom are calculated according to Satterthwaite approximation. In each study, we randomly sample 2,000 observations among all genes with replacement and then estimate the true parameters by different estimators and calculate corresponding losses. We replicate this study 2,000 times and calculate the risk by taking the average of the losses. (See Table 1.) The risk of  $\hat{\theta}_{SS}$  is about 92.8% of that of  $\hat{\theta}_{SM}$ , and 31.9% of that of  $\hat{\theta}_{NS}$ .

We have also calculated the standard deviation of the difference of the losses between an estimator  $\hat{\theta}$  and the double shrinkage estimator  $\hat{\theta}_{SS}$  and displayed it in the last row of Table 1. Clearly, the double shrinkage estimator improves shrinking-mean-alone estimator significantly which improves  $\hat{\theta}_{NS}$  significantly.

Along with the simulation studies we have presented in Section 4, we can state that the double shrinkage estimator  $\hat{\theta}_{SS}$  is better than the shrinking-mean-alone estimator  $\hat{\theta}_{SM}$ , which is better than the estimator  $\hat{\theta}_{NS}$  without any shrinkage.

The code for the double shrinkage estimator can be downloaded from <http://astro.temple.edu/~zhaozhg/publications.html>.



Table 1. The risk comparison of the estimators for golden spike-in data set. Within this table,

$DL = L(\hat{\theta}, \theta) - L(\hat{\theta}_{SS}, \theta)$ , the difference between the losses of any estimator  $\hat{\theta}$  and the double shrinkage estimator  $\hat{\theta}_{SS}$

Estimator	$\hat{\theta}_{NS}$	$\hat{\theta}_{SM}$	$\hat{\theta}_{SS}$
Risk	0.3243	0.1115	0.1035
E(DL)	0.2208	0.0080	0
Std(DL)	0.0107	0.0032	0

## 6. CONCLUSION AND DISCUSSION

In this article, we have constructed a new estimator when assuming the observation  $X_i$  follows a normal distribution with an unknown and unequal variance  $\sigma_i^2$ . The estimator is based on the model (6), a general form of *Log-Normal* model firstly proposed by Hwang and Liu (2010) and further studied in Hwang, Qiu and Zhao (2009). In these two papers, they have constructed the double shrinkage testing procedure and confidence interval by using the *empirical* Bayes approach. We adopt the *empirical* Bayes approach to construct a point estimator for multiple parameters which shrinks both the means and variances. We call this estimator  $\hat{\theta}_{SS}$  the double shrinkage estimator.

We further analyze the performance of  $\hat{\theta}_{SS}$ , comparing with the shrinking-mean-alone estimator  $\hat{\theta}_{SM}$  and the estimator  $\hat{\theta}_{NS} = X$  with no shrinkage. Both extensive simulation studies and a real data analysis indicate that  $\hat{\theta}_{SS}$  performs uniformly better than the other two. We thus strongly recommend the new approach.

This article proposes a new methodology in estimating under the condition of *heteroscedasticity*. However, much work is needed. For instance, we would like to know how  $\hat{\theta}_{SS}$  relates to the exact Bayes estimator in terms of the relative savings loss introduced in Efron and Morris (1973). However, the proof of any analytic results will be very difficult and heavily involved due to the unknown and unequal variances. We leave this for the future research.

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