

# Tail-summed scores method for inference in the Gaussian sequence model with applications to change-point analysis

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# Overview

- 1 TSS method
  - Choice of thresholds
- 2 Theoretical considerations
  - Perfect separation
  - Weak signal
- 3 Simulation results
- 4 Summary and extensions
- 5 Change-point applications

# The Gaussian sequence model

$$X_i = \mu_i + \varepsilon_i, \quad i = 1, \dots, n, \quad (1)$$

where  $\varepsilon_i \stackrel{iid}{\sim} N(0, \sigma^2)$  and  $\boldsymbol{\mu} = (\mu_1, \dots, \mu_n)$  is the unknown mean vector

- $\boldsymbol{\mu}$  is a sparse vector
- $\mathcal{S} = \{i : \mu_i \neq 0\}$ ,  $|\mathcal{S}| = k$
- $\sigma^2 = 1$

Inference on  $\boldsymbol{\mu}$ :

- Signal estimation
- Multiple testing

# TSS procedure

Sample:  $X_1, \dots, X_n$

Sequence of thresholds:  $\lambda_1, \dots, \lambda_n$

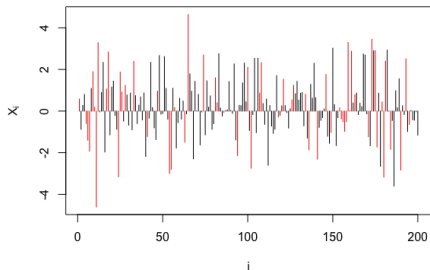
$$\textcircled{1} Y_i = X_i^2, i = 1, \dots, n$$

$$\textcircled{2} Y_{(n)} \geq \dots \geq Y_{(1)}$$

$$\textcircled{3} T_i = \sum_{j=1}^{n-i+1} Y_{(j)}$$

$$\textcircled{4} \hat{k} = \max_i \{T_i \geq \lambda_i\}$$

$$\hat{S} = \{j : Y_j \geq Y_{(n-\hat{k}+1)}\}$$



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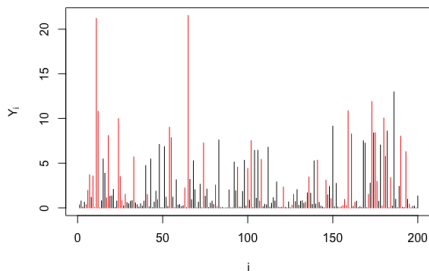
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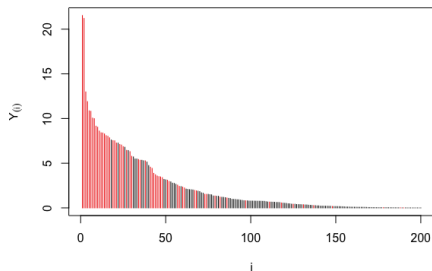
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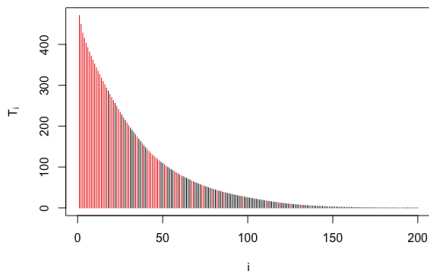
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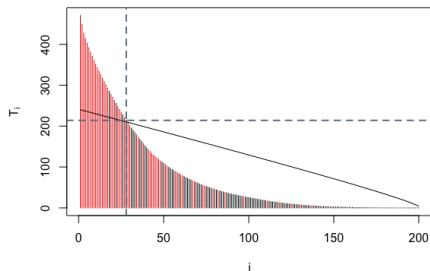
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## Choice of thresholds

Chi-square quantiles:

$$\lambda_i = q_{\chi_{n-i+1}^2}(1 - \alpha) \quad (2)$$

...or approximations:

$$\lambda_i^{H_i} = n - i + 1 + H_i \sqrt{2(n - i + 1)} \quad (3)$$

- $H_i = \sqrt{2 \log \frac{1}{\alpha}} + \sqrt{\frac{2}{n-i+1}} \log \frac{1}{\alpha}$
- $H_i = q_{N(0,1)}(1 - \alpha)$  or  $H_i = 2$
- $H_i = 0$  - asymptotically equal

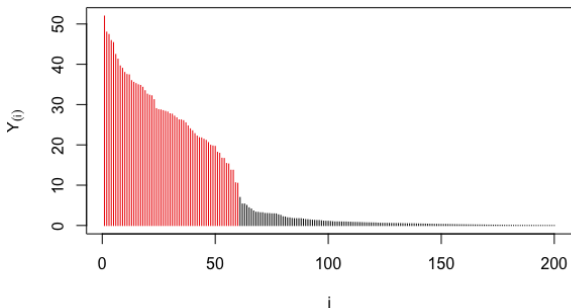
Contiguity property:

$$T_i \leq \lambda_i^{H_i} \implies T_{i+1} \leq \lambda_{i+1}^{H_{i+1}}$$

for any non-decreasing sequence  $H_i$ .

# Theoretical considerations - perfect separation

- [Duval et al., 2007] considered the TSS procedure with  $H_i = 0$
- Perfect separation event:  $\Omega_n = \{Y_{(1)}^S \geq Y_{(n-k)}^{NS}\}$
- Perfect separation assumption:  $P(\Omega_n) \rightarrow 1, n \rightarrow \infty$
- A sufficient condition:  $\mu \geq \sqrt{2 \log(n-k)} + \sqrt{2 \log k}$



# Perfect separation

Let

- $k = \pi_1 n$  for some  $\pi_1 \in (0, 1)$
- The perfect separation assumption holds
- $\lambda_i^{H_i}$  such that  $H_i(n)/\sqrt{nu_n} \rightarrow 0$ , where  $u_n$  is s.t.  $u_n \rightarrow 0$ ,  $\sqrt{nu_n} \rightarrow \infty$

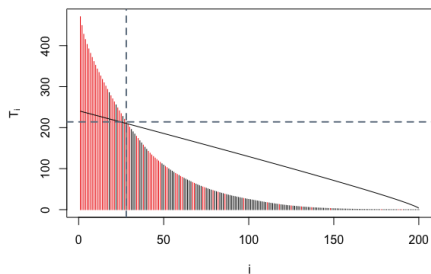
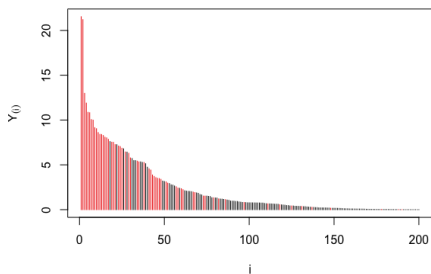
For the TSS stopping time  $\hat{k}$  it holds that

$$P \left( \left| \frac{\hat{k}}{n} - \pi_1 \right| \geq u_n \right) \rightarrow 0, \quad n \rightarrow \infty$$

$$FDR \rightarrow 0$$

$$FNR \rightarrow 0$$

# Weak signal - no assumptions on the strength



# Weak signal - no assumptions on the strength

- **Stop after the mixing starts**

Let  $j$  be the position of the largest non-signal variable in the decreasingly sorted sample. It holds that

$$P\left(\hat{k} \leq j\right) \rightarrow 0, \quad n \rightarrow \infty.$$

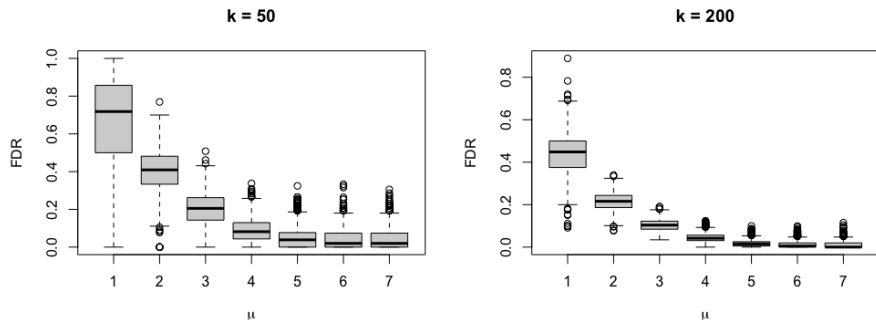
- **Stop before the mixing ends**

The probability of the TSS method overestimating the number of signals  $k$  is small:

$$P\left(\hat{k} > k\right) \leq 1 - F_{\chi_{n-k}^2}(\lambda_{k+1}).$$

# Simulation results

The TSS procedure does not control the FDR when the signal is weak.



**Figure:** Boxplots of the estimated FDR for the TSS procedure with  $H = 0$ ,  $n = 1000$ .

# Simulation results

The TSS procedure works well as a signal estimation method if the signal is weak.

Number nonzero	$\mu = 2$					$\mu = 3$				
	50	100	200	300	500	50	100	200	300	500
<b>TSS</b> $H = 2$	<b>206</b>	<b>377</b>	<b>662</b>	904	1300	275	434	689	888	1190
<b>TSS</b> $H = 0$	209	<b>367</b>	<b>634</b>	<b>859</b>	<b>1226</b>	<b>227</b>	374	609	798	1078
UNI	<b>203</b>	404	805	1206	2000	366	725	1433	2122	3391
FDR $q = 0.05$	314	383	785	1150	1795	313	519	836	1085	1460
FDR $q = 0.2$	227	421	673	900	1230	<b>226</b>	<b>350</b>	<b>527</b>	<b>659</b>	<b>850</b>
EBT	<b>203</b>	386	681	<b>874</b>	<b>944</b>	242	<b>361</b>	<b>494</b>	<b>585</b>	<b>749</b>
SURE	<b>203</b>	400	803	1206	2012	312	700	1443	2180	3640

**Table:** The estimated  $l_2$  risk of different thresholding estimators based on  $N = 1000$  repetitions for the sample of size  $n = 1000$  from the Gaussian sequence model.

# Summary and extensions

- Proposed is a new thresholding signal estimation/Multiple testing procedure for the Gaussian sequence model
  - Consider values in groups
  - Stop when the remaining signal is too weak
- Choose the sequence of thresholds depending on the objective:
  - FWER control
  - Signal estimation
  - FDR control
  - Estimating the  $l_2$  norm of the quadratic functional [Collier et al., 2017]

$$\hat{\mu}_{\text{minimax}} = \sum_{i=1}^n X_i^2 - n$$

- Different transformation of the data?



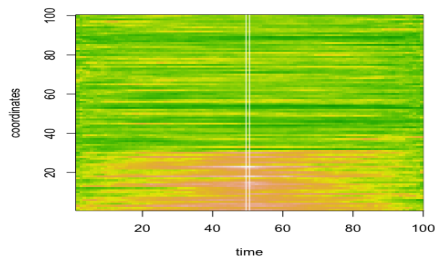
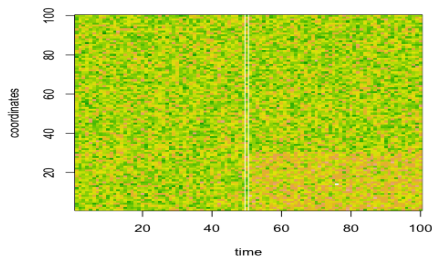
# Change-point applications

$$X_{ij} = \mu_{ij} + \varepsilon_{ij}, \quad i = 1, \dots, p \quad j = 1, \dots, n,$$

$$\mathcal{S} = \{j : \mu_{i,\tau+1} \neq \mu_{i,\tau}\} \subset \{1, \dots, p\}.$$

CUSUM transform the data matrix  $X \in \mathbb{R}^{p \times n} \mapsto Z \in \mathbb{R}^{p \times (n-1)}$  is

$$Z_{ij} = \sqrt{\frac{j(n-j)}{n}} \left( \frac{1}{n-j} \sum_{l=j+1}^n X_{il} - \frac{1}{j} \sum_{l=1}^j X_{il} \right)$$



# Change-point applications

$$Z_{ij} \sim N(\theta_{ij}, 1),$$

$$Z_{\cdot, \tau} \sim N(\theta_{\cdot, \tau}, I_d), \text{ where } \theta_{\cdot, \tau} = \sqrt{\frac{\tau(n-\tau)}{n}}(\mu_{\cdot, \tau+1} - \mu_{\cdot, \tau})$$

This separates the coordinates into two groups based on the distribution of  $Z_{i, \tau}$ :

$$Z_{i, \tau} \sim \begin{cases} N(0, 1), & i \notin \mathcal{S} \\ N(\theta_{i, \tau}, 1), & i \in \mathcal{S} \end{cases} \quad (4)$$

# Change-point applications - inference on $\mathcal{S}$

Consistent estimation of  $\mathcal{S}$

- [Jirak, 2015]

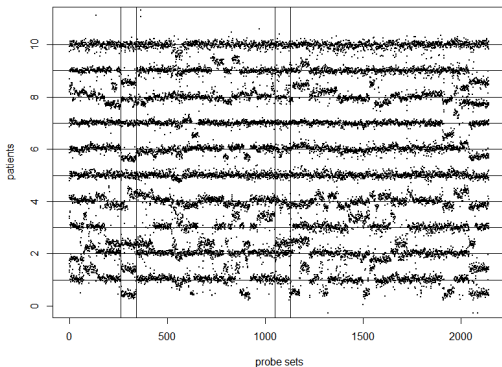
As a part of the change-point estimation algorithm

- [Cho and Fryzlewicz, 2015], [Cho, 2016]
- [Wang and Samworth, 2018]

Higher Criticism-like statistic for change-point detection

- [Jeng et al., 2012]

# Copy number variation (CNV) data



**Figure:** Log-intensity-ratio values for 10 individuals at 2215 different loci from the study of 43 patients with a bladder tumour. Positive values indicate duplication, while negative indicate deletion at a given locus.

- Variants can be shared among a large number of patients or individual-specific

# Change-point applications - inference on $\mathcal{S}$

- Estimating  $\mathcal{S}$ 
  - Improving the accuracy of a change-point estimator
  - For further research on the affected profiles

Multiple testing or signal estimation methods

- Estimating  $|\mathcal{S}|$ 
  - Estimating the sparsity of the problem
  - Estimating the frequency of copy number variants across a population to get a measure of the extent of the changes studied

Methods for estimating the proportion of the false null hypotheses in multiple testing

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Collier, O., Comminges, L., and Tsybakov, A. B. (2017).

Minimax estimation of linear and quadratic functionals on sparsity classes.  
*The Annals of Statistics*, 45(3):923–958.

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-  Duval, M., Delmas, C., Laurent, B., and Robert-Granié, C. C. (2007). A procedure based on partial sums of order statistics to detect differentially expressed genes. Working paper or preprint, available at [https://hal.archives-ouvertes.fr/hal-00302355/file/Duval\\_etal\\_arxiv.pdf](https://hal.archives-ouvertes.fr/hal-00302355/file/Duval_etal_arxiv.pdf).
-  Jeng, X. J., Cai, T. T., and Li, H. (2012). Simultaneous discovery of rare and common segment variants. *Biometrika*, 100(1):157–172.
-  Jirak, M. (2015). Uniform change point tests in high dimension. *The Annals of Statistics*, 43(6):2451–2483.

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