

# Rate-optimal gamma scale mixture detection

Joint work with Qikun Chen

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# Outline

Motivating example: ion channel openings

Gamma scale mixture detection

The case  $0 < \theta < 1$  for general  $\alpha$ .

**Motivating example: ion channel openings**

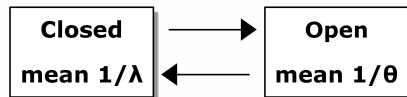
## Modelling ion channels opening times

- Neurotransmitter is released across the synapse via the opening and closing of calcium ion channels.
- Opening times can be measured.
- One scientific question has been:

**Is there a single open state or multiple open states?**

## Continuous-time Markov chains

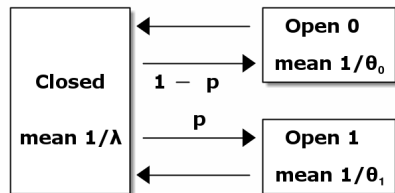
- Times in different states have been modelled using continuous-time Markov chains.
  - Times in each state are exponentially distributed.
- If only a single “open” state, a series of measurements should resemble a sample from an exponential distribution.
- Each observed opening time  $X_i$  satisfies  $P(X_i > x) = e^{-\theta x}$ .



## Multiple states give a mixture

- If there are two (different) *open* states but the measuring device cannot distinguish between them, opening times form a **mixture** of two exponential samples.
- Each observed opening time  $X_i$  satisfies

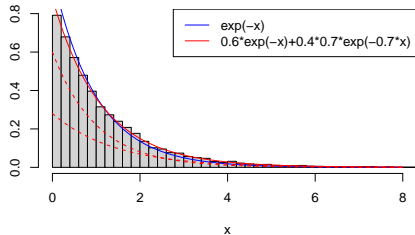
$$P(X_i > x) = (1 - p)e^{-\theta_0 x} + pe^{-\theta_1 x}.$$



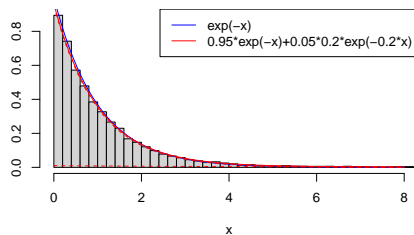
## Hard-to-detect local alternatives

- Choosing between one or two open states can be formulated as a hypothesis-testing problem.
- There are two ways a “local alternative” can be hard to detect:
  1. the two means are close to each other
  2. the mixing proportion  $p$  is close to 0 (or 1)
- These two have quite different behaviour, case 2. being the “most challenging”.

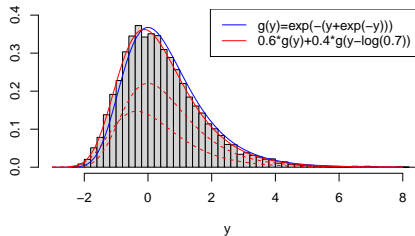
**X ~ Exponential scale mixture**



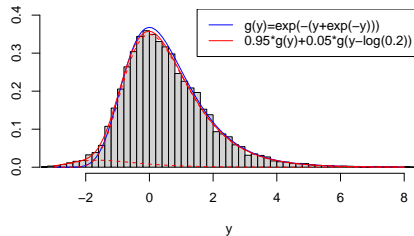
**X ~ Exponential scale mixture**



**Y = -log(X) ~ Gumbel location mixture**



**Y = -log(X) ~ Gumbel location mixture**





## **Gamma scale mixture detection**

## Hypothesis testing problem of interest

- We model data as iid random variables  $X_1, \dots, X_n$ .
- Let  $F_\alpha(\cdot)$  denote the gamma( $\alpha, 1$ ) CDF ( $\alpha$  is **known**).
- We are interested in the hypothesis testing problem

$$H_0: P(X_1 \leq x) = F_\alpha(x) \text{ vs. } H_1: P(X_1 \leq x) = (1 - p)F_\alpha(x) + pF_\alpha(\theta x),$$

for  $0 < p < 1$  and  $\theta \neq 1$ .

- This is the simplest possible “gamma scale mixture detection” model.

# Distinguishability

- For a given sequence  $(p_n, \theta_n)$ , write  $G_n(x) = (1 - p_n)F_\alpha(x) + p_nF_\alpha(\theta_n x)$ .
- We address the question:

**How close to  $F_\alpha(\cdot)$  can the mixture  $G_n(\cdot)$  be and still be “detectable”?**

That is, is there a test so that under  $G_n(\cdot)$  we have power  $\rightarrow 1$ ?

- We focus on the “**sparse mixture**” case where  $p_n \rightarrow 0$ .

## Previous work

- Deep general results for exponential families provided by Ditzhaus (2019) covered the gamma scale mixture where  $\theta > 1$ ;
  - this is when the contaminating mean is **smaller** than the null.
- Arias-Castro and Huang (2020) covered the case  $0 < \theta < 1$  for  $\alpha = \frac{1}{2}$ ;
  - this is a  $\chi_1^2$  scale mixture;
  - applicable if we have a normal mixture variance with common known mean.

**The case  $0 < \theta < 1$  for general  $\alpha$ .**

## Separation in $n$ dimensions

- In one sense the problem is “easy”: if
  - $F_\alpha^n$  and  $G_n^n$  are the  $n$ -dimensional versions of  $H_0$  and  $H_1$ ;
  - if the **total variation** (TV, i.e. half  $L_1$ ) distance

$$d_{\text{TV}}(F_\alpha^n, G_n^n) = \sup_A |F_\alpha^n\{A\} - G_n^n\{A\}| \rightarrow \begin{cases} 1 & \text{then NP test has limiting power 1;} \\ 0 & \text{then NP test has no limiting power.} \end{cases}$$

- The difficulty is in approximating TV distance in  $n$  dimensions.

## Hellinger distance trick

- The Hellinger distance whose square is

$$d_H^2(F_\alpha^n, G_n^n) = \int \left( \sqrt{dF_\alpha^n} - \sqrt{dG_n^n} \right)^2 = 2 \left[ 1 - \int \sqrt{dF_\alpha^n dG_n^n} \right]$$

$\rightarrow 0$  if and only if  $d_{TV}(F_\alpha^n, G_n^n) \rightarrow 0$ .

- Since  $\int \sqrt{dF_\alpha^n dG_n^n} = \left( \int \sqrt{dF_\alpha dG_n} \right)^n$  there is a nice relationship between  $d_H(F_\alpha, G_n)$  and  $d_H(F_\alpha^n, G_n^n)$ :

$$d_H^2(F_\alpha^n, G_n^n) = 2 \left\{ 1 - \left[ 1 - \frac{1}{2} d_H^2(F_\alpha, G_n) \right]^n \right\}.$$

- So if  $nd_H^2(F_\alpha, G_n) \rightarrow 0$ ,  $d_{TV}(F_\alpha^n, G_n^n) \rightarrow 0$  too.

## Critical rate $r_n$ under 4 scenarios

- In S. (2022) we showed that under each scenario for  $\theta_n$ , that if  $p_n = o(r_n)$ ,  $nd_H^2(F_\alpha, G_n) \rightarrow 0$ :

	Scenario	$r_n$
1.	$\theta_n = 1 - \Delta_n \uparrow 1$	$n^{-1/2} \Delta_n^{-1}$
2.	$\theta_n \equiv \theta \in (\frac{1}{2}, 1)$ fixed	$n^{-1/2}$
3.	$\theta_n \equiv \frac{1}{2}$ fixed	$[n(\log n)^\alpha]^{-1/2}$
4.	$\theta_n \equiv \theta \in (0, \frac{1}{2})$ fixed	$[n(\log n)^{\alpha-1}]^{\theta-1}$

► slide 15



## Bonferroni test attains $r_n$ in 3 scenarios

- A test based on the sample mean or median can detect  $p_n \asymp r_n$  (i.e. attains critical rate) in scenarios 1 and 2.
- A test based on the sample maximum attains the critical rate in scenario 4.
- A Bonferroni test using the smallest p-value of these two attains the critical rate in scenarios 1, 2 and 4;
  - it does **not** attain the critical rate in scenario 3 ( $\theta_n \equiv \frac{1}{2}$ ) though.

## Score test attains critical rate in scenario 3

- The score statistic for testing  $H_0: p = 0$  vs.  $H_1: p > 0$  when  $\theta \equiv \frac{1}{2}$  is known is  $\sum_{i=1}^n e^{X_i/2}$ .
- Note that under  $H_0$ ,  $\text{Var}(\sum_{i=1}^n e^{X_i/2}) = \infty$ !
- Nonetheless, in Chen and S. (2024) we showed that under scenario 3,

$$\frac{\sum_{i=1}^n [e^{X_i/2} - 2^\alpha]}{\sqrt{n(\log n)^\alpha}} \xrightarrow{d} \begin{cases} N(0, \Gamma(\alpha + 1)^{-1}) & \text{if } p_n \equiv 0, \\ N(\mu, \Gamma(\alpha + 1)^{-1}) & \text{if } p_n \sim \frac{\mu 2^\alpha \Gamma(\alpha + 1)}{\sqrt{n(\log n)^\alpha}} \end{cases}$$

and thus attains the critical rate.

- Thus a Bonferroni test based on the smallest of 3 p-values (this score test plus sample mean and max) attains critical rate in all 4 scenarios.

## Current/future work

- Note that the power of  $\log n$  in  $r_n$  varies in a non-continuous way as  $\theta \rightarrow \frac{1}{2}$ .

◀ slide 12

- It turns out that in a scenario where  $\theta_n \rightarrow \frac{1}{2}$  *slowly enough* then a different critical rate is obtained:
  - e.g. if  $\theta_n = \frac{1+\Delta_n}{2} \downarrow \frac{1}{2}$ ,  $r_n = \sqrt{\frac{\max\left(\Delta_n, \frac{1}{\log n}\right)^\alpha}{n}}$
  - the Bonferroni test may not attain this critical rate;
  - indeed *no* adaptive test (i.e. without knowledge of  $\Delta_n$ ) may be able to.
- A broader “asymptotic minimax” framework may be needed:
  - a “price for adaptivity” like the extra  $\sqrt{\log \log n}$  factor seen in the analogous normal location mixture problem (see Ingster (1997, 2001, 2002)), may apply;
  - the GLRT would then be optimal according to that, and thus **not** attain the critical rates in our 4 scenarios.

**THANK YOU!**

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