

Mathematical Statistics II

STA2212H S LEC9101

Week 6

February 11 2025

School phone bans alone do not improve grades or wellbeing, says UK study

Researchers say bans need to be part of wider strategy to tackle negative impact of mobile use on children



The negative effects of phone overuse did not differ between schools that banned phones and those that did not.

School phone policies and their association with mental wellbeing, phone use, and social media use (SMART Schools): a cross-sectional observational study

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Today

1. Recap Feb 4 decision theory, efficiency
2. Theory of interval estimation
3. Approximations to the posterior
4. Project: Choice of papers ↪
5. HW5, Statistics in the News

MS 7.1

MS 7.2

The slide is a promotional graphic for a Statistics Colloquium. It features a dark blue header with the University of Toronto Statistical Sciences logo. Below the header is a photo of a woman with dark hair, identified as Pratheepa Jeganathan. To her left is her name in large white text. Below her name is a brief bio: "Assistant Professor in the Department of Mathematics and Statistics, McMaster University". A green bar on the left contains the text "UPCOMING SPEAKER". On the right, a white box displays the date "13 Feb" and time "11:00 am room 9014". The main title "STATISTICS COLLOQUIUM" is at the bottom in white, along with a subtitle: "A Robust Nonparametric Framework for Detecting Repeated Spatial Clusters". A detailed abstract of the talk follows.

Statistical Sciences
UNIVERSITY OF TORONTO

PRATHEEPA JEGANATHAN

Assistant Professor in the Department of Mathematics and Statistics, McMaster University

UPCOMING SPEAKER

STATISTICS COLLOQUIUM

A Robust Nonparametric Framework for Detecting Repeated Spatial Clusters

Ensuring spatial contiguity in spatial clustering is often critical, as nearby observations exhibit dependencies. Equally important, however, is identifying repeated spatial clusters that may not be spatially contiguous. Traditional clustering techniques such as constrained hierarchical clustering, constrained partitioning, and density-based clustering, are commonly used to ensure spatial contiguity. Despite their strengths, these methods often fail to detect repeated spatial patterns consistently, particularly under varying degrees of spatial dependence. This talk introduces a post-clustering framework to enhance constrained clustering techniques by identifying repeated spatial clusters. Specifically, constrained clustering methods often overestimate the number of clusters when repeated patterns are present. Our framework incorporates a nonparametric test based on Maximum Mean Discrepancy (MMD) and block permutation to assess the distributions of multivariate attributes within the clusters identified by constrained methods. I will discuss the performance of the proposed framework through a simulation study, evaluating its robustness across varying levels of spatial dependence, cluster shapes, the number of multivariate attributes, and spatial locations.

13 Feb
11:00 am
room 9014

700 University Ave. 9th Floor
Department of Statistical Sciences
University of Toronto

statistics.utoronto.ca

Department Seminar Thursday February 13 11.00 – 12.00

Hydro Building, Room 9014

Robust nonparametrics and spatial data

Pratheepa Jeganathan, McMaster University

Recap: Decision theory

AoS Ch 12, MS Ch 6.2

- Loss function

$$L(\hat{\theta}, \theta)$$

measures "dist" from $\hat{\theta}$ to θ

squared-error, absolute error, ...

- Risk function

exp'd loss

$$\int L(\hat{\theta}(x), \theta) f(x; \theta) d\theta = R_{\theta}(\hat{\theta})$$

expected loss

- Admissible point estimators

no other est. with an
always better risk f.

not inadmissible

$$R_{\theta}(\hat{\theta}) > R_{\theta}(\tilde{\theta})$$

average over $\pi(\theta)$

- Bayes risk

$$R_B(\hat{\theta}) = \int R_{\theta}(\hat{\theta}) \pi(\theta) d\theta$$

minimize Bayes risk

- Bayes estimator

$\hat{\theta}$
minimizes Bayes risk

> for some $\underline{\theta}$

Optimal Bayes estimators

MS 6.2

- Bayes risk

$$\min_{\hat{\theta}} R_B(\hat{\theta}) = \int R_\theta(\hat{\theta}) \pi(\theta) d\theta = \int L(\hat{\theta}(x), \theta) f(x; \theta) dx \pi(\theta) d\theta$$

- Optimal Bayes estimators minimize the Bayes risk

- Equivalent to minimizing posterior loss: $\int L\{\hat{\theta}(x), \theta\} \pi(\theta | x) d\theta$

$$= \min_{\hat{\theta}} \int \int L(\hat{\theta}(x|\theta) \pi(\theta|x) d\theta \frac{m(x)}{dx}$$

- Example: absolute-error loss

$$L(\hat{\theta}, \theta) = |\hat{\theta} - \theta|$$

achieved at $\hat{\theta} = \bar{x}$

$$\min_{\hat{\theta}} \int |\hat{\theta} - \theta| \pi(\theta | x) d\theta$$

- solution $\hat{\theta}(x) = \text{median } \pi(\theta | x)$

$$L(\hat{\theta}, \theta) = (\hat{\theta} - \theta)^2$$

$E(\theta | x)$ post. mean.

posterior median

- Suppose $\hat{\theta}$ is a Bayes estimator and is unique

requires proper prior

- Suppose we have another estimator $\tilde{\theta}$ with a smaller frequentist risk function:

$$R_{\theta}(\tilde{\theta}) \leq R_{\theta}(\hat{\theta})$$

- The Bayes risk of $\tilde{\theta}$ is $R_B(\tilde{\theta}) = \int \underbrace{R_{\theta}(\tilde{\theta})}_{\text{ }} \pi(\theta) d\theta$

$$\leq \int R_{\theta}(\hat{\theta}) \pi(\theta) d\theta = R_B(\hat{\theta}) \quad \text{contradiction}$$

- Suppose $\hat{\theta}$ is a Bayes estimator and is unique
- Suppose we have another estimator $\tilde{\theta}$ with a smaller frequentist risk function:

$$R_\theta(\tilde{\theta}) \leq R_\theta(\hat{\theta})$$

- The Bayes risk of $\tilde{\theta}$ is $R_B(\tilde{\theta}) = \int$
- instead of finding estimator to minimize the weighted average of the risk function we could

$$\min_{\hat{\theta}} \max_{\theta} R_\theta(\hat{\theta})$$

- such estimators are called **minimax** (not quite admissible)
- Bayes estimators with constant risk are minimax

Definition §6.2
admissible

$$X \sim \text{Bin}(n, \theta)$$

$$\text{Beta}(\alpha, \beta) \quad \alpha = \beta = \frac{1}{n}$$

- finding the ‘best’ point estimator $\hat{\theta}$
- best = smallest expected loss
- no asymptotic theory involved
- can find these using a Bayesian argument
- but the justification is not Bayesian
- another non-asymptotic approach to ‘best’ estimators: UMVUE

unbiased
=

MS 6.3

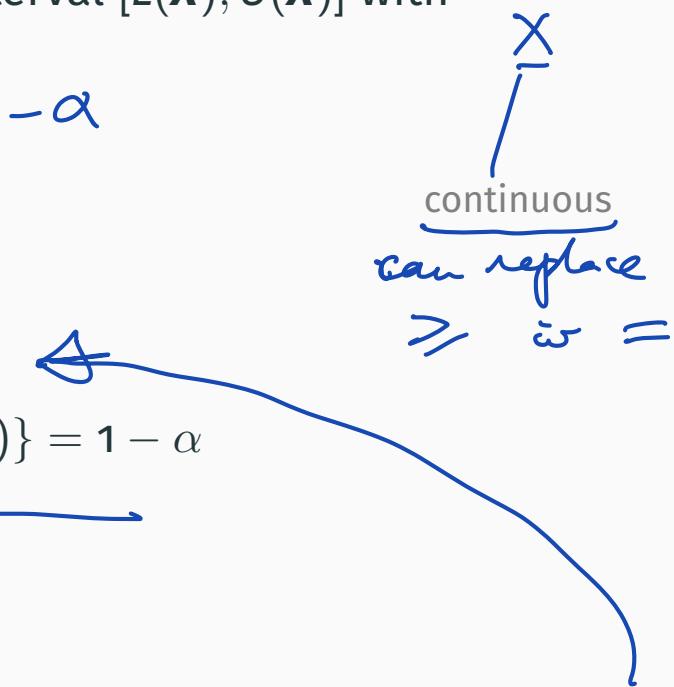
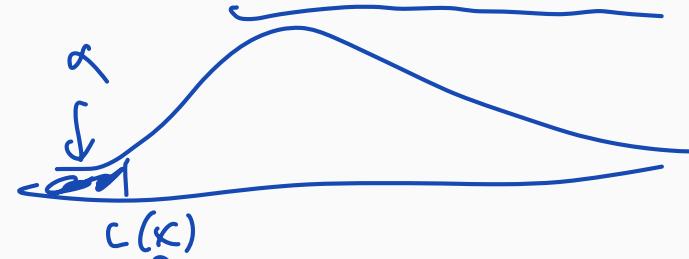
- X_1, \dots, X_n i.i.d. $f(x; \theta), \theta \in \mathbb{R}$
- a $100(1 - \alpha)\%$ confidence interval for θ is a random interval $[L(\mathbf{X}), U(\mathbf{X})]$ with

$$\Pr_{\theta} \{ L(\mathbf{X}) \leq \theta \leq U(\mathbf{X}) \} \geq 1 - \alpha$$

↑
under $f(\cdot; \theta)$

- similarly, upper and lower $(1 - \alpha)$ -confidence bounds:

$$\Pr\{\theta \geq L(\mathbf{X})\} = 1 - \alpha; \quad \Pr\{\theta \leq U(\mathbf{X})\} = 1 - \alpha$$



recall confidence dist

- X_1, \dots, X_n i.i.d. $f(x; \theta), \theta \in \mathbb{R}$
- a $100(1 - \alpha)\%$ confidence interval for θ is a random interval $[L(\mathbf{X}), U(\mathbf{X})]$ with

continuous

- similarly, upper and lower $(1 - \alpha)$ -confidence bounds:

$$\text{pr}\{\theta \geq L(\mathbf{X})\} = 1 - \alpha; \quad \text{pr}\{\theta \leq U(\mathbf{X})\} = 1 - \alpha$$

- exact limits if we have exact distribution of \mathbf{X}
- approximate limits if $\text{pr}_{\theta}\{L(\mathbf{X}) \leq \theta \leq U(\mathbf{X})\} \approx 1 - \alpha$

- Example: X_1, \dots, X_n i.i.d. $N(\mu, 1)$

pivotal quantity = $\{g(\bar{x}; \theta) \text{ w property that } g(\bar{x}; \theta) \text{ is fixed } \forall \theta\}$

$$\bar{X} \sim N(\mu, \frac{1}{n}) \quad (\bar{X} - \mu) \sim N(0, \frac{1}{n})$$

$$\sqrt{n}(\bar{X} - \mu) \sim N(0, 1)$$

$$\Pr[-1.96 \leq Z \leq 1.96] = 0.95 \quad Z \sim N(0, 1)$$

$$= \Pr[-1.96 \leq \sqrt{n}(\bar{X} - \mu) \leq 1.96]$$

$$= \Pr\left\{ \bar{X} - \frac{1.96}{\sqrt{n}} \leq \mu \leq \bar{X} + \frac{1.96}{\sqrt{n}} \right\}$$

Exact confidence intervals

MS 7.1

- Example: X_1, \dots, X_n i.i.d. $N(\mu, 1)$
- Example X_1, \dots, X_n i.i.d. $U(0, \theta)$

$$\hat{\theta} = X_{(n)}$$

$$n(X_{(n)} - \theta) \xrightarrow{d} \exp(1)$$

$\frac{X_{(n)}}{\theta}$ has known dist: $\Pr(a \leq \frac{X_{(n)}}{\theta} \leq b)$

$$f(x_i) = \frac{1}{\theta}, 0 \leq x_i \leq \theta$$

$$F(x_i) = \frac{x_i}{\theta}, 0 \leq x_i \leq \theta$$

$$F_{X_{(n)}}(x) = \Pr(X_{(n)} \leq x) = \prod_{i=1}^n \left(\frac{x}{\theta}\right) = \left(\frac{x}{\theta}\right)^n$$

$$\Pr(a \leq \frac{X_{(n)}}{\theta} \leq b) = b^n - a^n$$

$$= \Pr(bX_{(n)} \leq \theta \leq aX_{(n)})$$

choose a, b s.t. $b^n - a^n = 1 - \alpha$

$$z = \frac{X_{(n)}}{\theta}$$

$$F_z(z) = z^n, 0 \leq z \leq 1$$

Exact confidence intervals

minimize $(b-a) \cdot X_{(n)}$ s.t. a, b

MS 7.1

• It can be shown that

$$b = 1, a = (1-\alpha)^{1/n} \text{ "pr}\left(a \leq \frac{X_{(n)}}{\theta} \leq b\right)$$

- Example: X_1, \dots, X_n i.i.d. $N(\mu, 1)$
- Example X_1, \dots, X_n i.i.d. $U(0, \theta)$
- Example $\underline{X_1, \dots, X_n}$ i.i.d. $N(\mu, \sigma^2)$

$$P_n \left\{ \frac{(\bar{X} - \mu)}{S/\sqrt{n}} \leq t \right\} = P_n \left\{ T_{n-1} \leq t \right\} \xrightarrow{\text{student's t-distr}}$$

$$ES^2 = \sigma^2 \quad S^2 = \frac{1}{n-1} \sum (X_i - \bar{X})^2 \quad \bar{X} = \frac{1}{n} \sum X_i$$

* $\rightarrow P_n \left\{ -t_{n-1}^{1/2} \leq \frac{\sqrt{n}(\bar{X} - \mu)}{S} \leq t_{n-1}^{1/2} \right\} = 1-\alpha \quad \text{by def}$

$$= P_n \left(\bar{X} - t_{n-1} \frac{S}{\sqrt{n}} \leq \mu \leq \bar{X} + t_{n-1} \frac{S}{\sqrt{n}} \right)$$

95% values
?

Approximate confidence intervals

MS 7.1

- Example: $X \sim \text{Binom}(n, \theta)$, $\hat{\theta} \sim N(\theta, \theta(1-\theta)/n)$

MS Ex.7.6

$$\Pr_{\theta} \left[-1.96 \leq \frac{\sqrt{n}(\hat{\theta} - \theta)}{\{\theta(1-\theta)\}^{1/2}} \leq 1.96 \right] \approx 0.95$$

$$\hat{\theta} - \theta \sim N(0, \frac{\theta(1-\theta)}{n})$$

$$\sqrt{n} \left(\frac{\hat{\theta} - \theta}{\sqrt{\theta(1-\theta)}} \right) \sim N(0, 1)$$

approximate first

 $I_n^{-1}(\theta)$ as
approx variance
of $\hat{\theta}$ solve for θ means solve a quadratic eq =

$$\frac{n(\hat{\theta} - \theta)^2}{\theta(1-\theta)} \leq 1.96$$

$$n(\hat{\theta}^2 - 2\hat{\theta}\theta + \theta^2) \leq 1.96\theta - 1.96\theta^2$$

Approximate confidence intervals

MS 7.1

- Example: $X \sim \text{Binom}(n, \theta)$, $\hat{\theta} \sim N(\theta, \theta(1-\theta)/n)$

MS Ex.7.6

$$\text{pr}_{\theta} \left[-1.96 \leq \frac{\sqrt{n}(\hat{\theta} - \theta)}{\{\theta(1-\theta)\}^{1/2}} \leq 1.96 \right] \approx 0.95$$

$$\text{pr}_{\theta} \left[-1.96 \leq \frac{\sqrt{n}(\hat{\theta} - \theta)}{\{\hat{\theta}(1-\hat{\theta})\}^{1/2}} \leq 1.96 \right] \approx 0.95$$

$$\uparrow I_n(\hat{\theta})$$

pt. of

$$\hat{\theta} \pm 1.96 \frac{\hat{\theta}(1-\hat{\theta})}{\sqrt{n}}$$

dividing by $\hat{\theta}$ is called
standardizing

- Example: $X \sim \text{Binom}(n, \theta)$, $\hat{\theta} \sim N(\theta, \theta(1-\theta)/n)$

MS Ex.7.6

$$\text{pr}_{\theta} \left[-1.96 \leq \frac{\sqrt{n}(\hat{\theta} - \theta)}{\{\theta(1-\theta)\}^{1/2}} \leq 1.96 \right] \approx 0.95$$

$$\text{pr}_{\theta} \left[-1.96 \leq \frac{\sqrt{n}(\hat{\theta} - \theta)}{\{\hat{\theta}(1-\hat{\theta})\}^{1/2}} \leq 1.96 \right] \approx 0.95$$

- $\hat{\theta}_n$ maximum likelihood estimate
- approximate 95% confidence interval

$$\hat{\theta} \sim N[\theta, \{nl(\theta)\}^{-1}]$$

AoS Thm 6.16

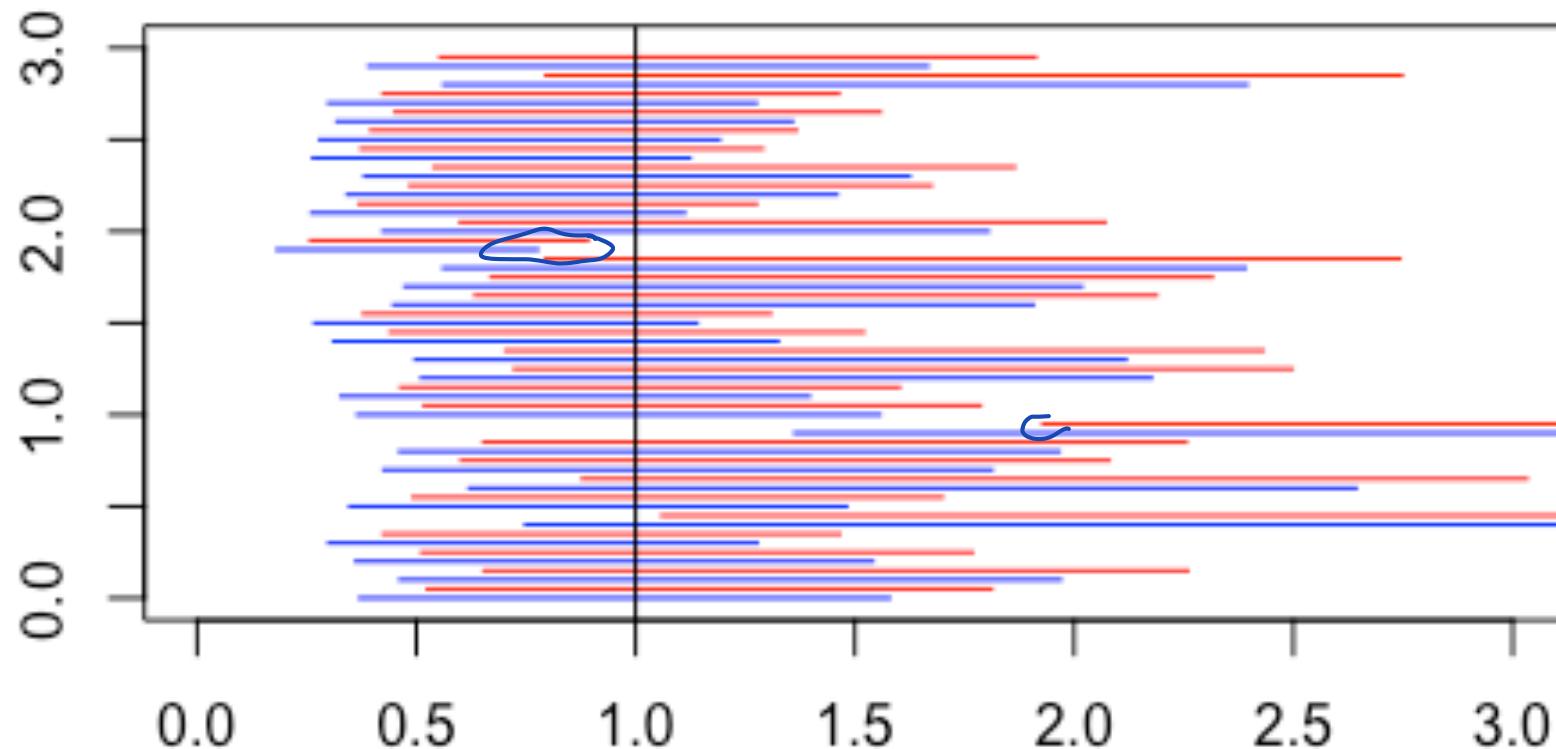
$$\hat{\theta} \pm z_{\alpha/2} \hat{s}\hat{e}$$

- X_1, \dots, X_n i.i.d. $\text{Exp}(\lambda)$ $f(x_i; \lambda) = \lambda e^{-\lambda x_i}$; $x_i > 0, \lambda > 0$

- $\hat{\lambda} = \underline{\underline{\frac{1}{\bar{x}}}}$

$$\beta \leftarrow g(\lambda) = \log \lambda \quad \text{gfg} \quad \hat{\phi} = g(\hat{\lambda}) = -\log \bar{x}$$

n= 10



- upper and lower bounds

$$\theta \in \mathbb{R}$$

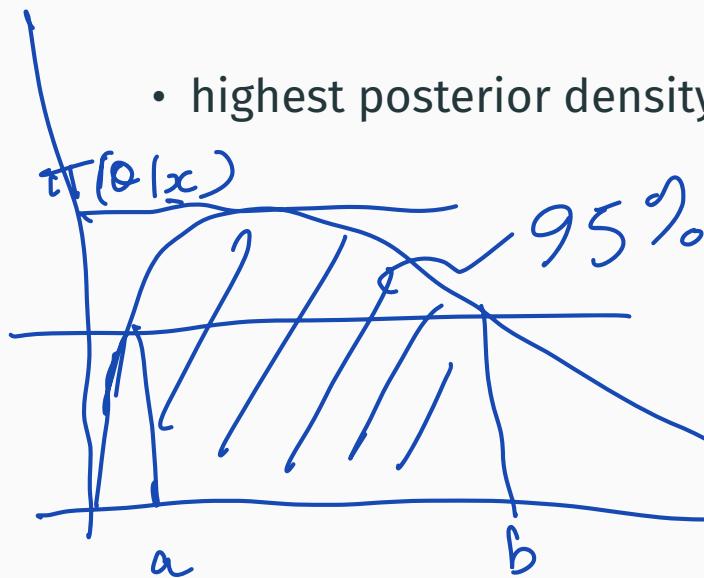
- upper and lower bounds $\theta \in \mathbb{R}$
- equi-tailed posterior intervals

- upper and lower bounds $\theta \in \mathbb{R}$
- equi-tailed posterior intervals
- highest posterior density $\theta \in \mathbb{R}^p, p \geq 1$

... Bayesian credible intervals

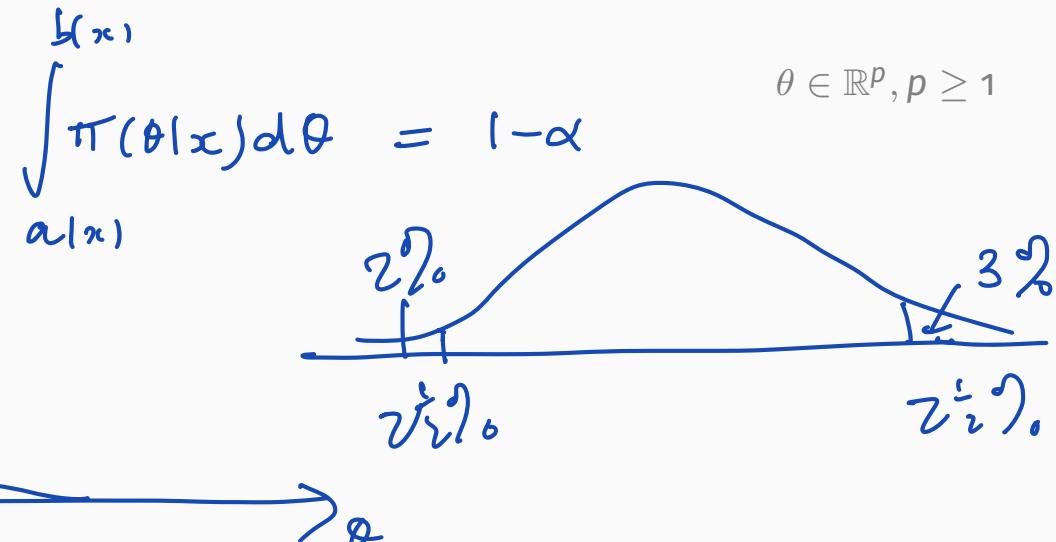
MS 7.2

- upper and lower bounds
- equi-tailed posterior intervals



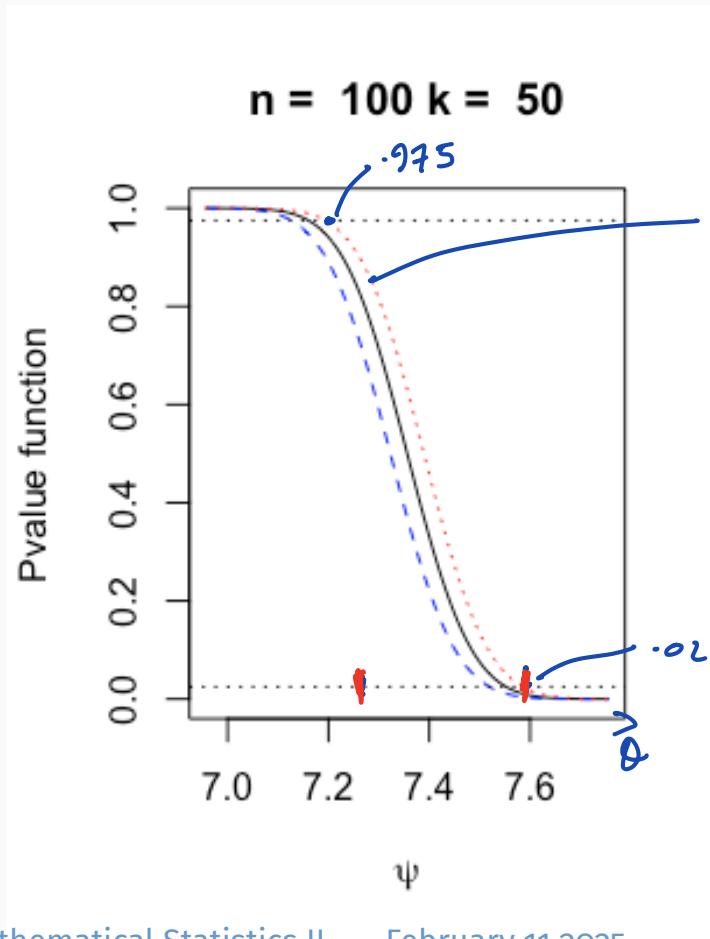
$\pi(\theta|x)$ gives all our probabilities

$$\theta \in \mathbb{R}$$



Example: Equi-tailed Bayesian credible intervals

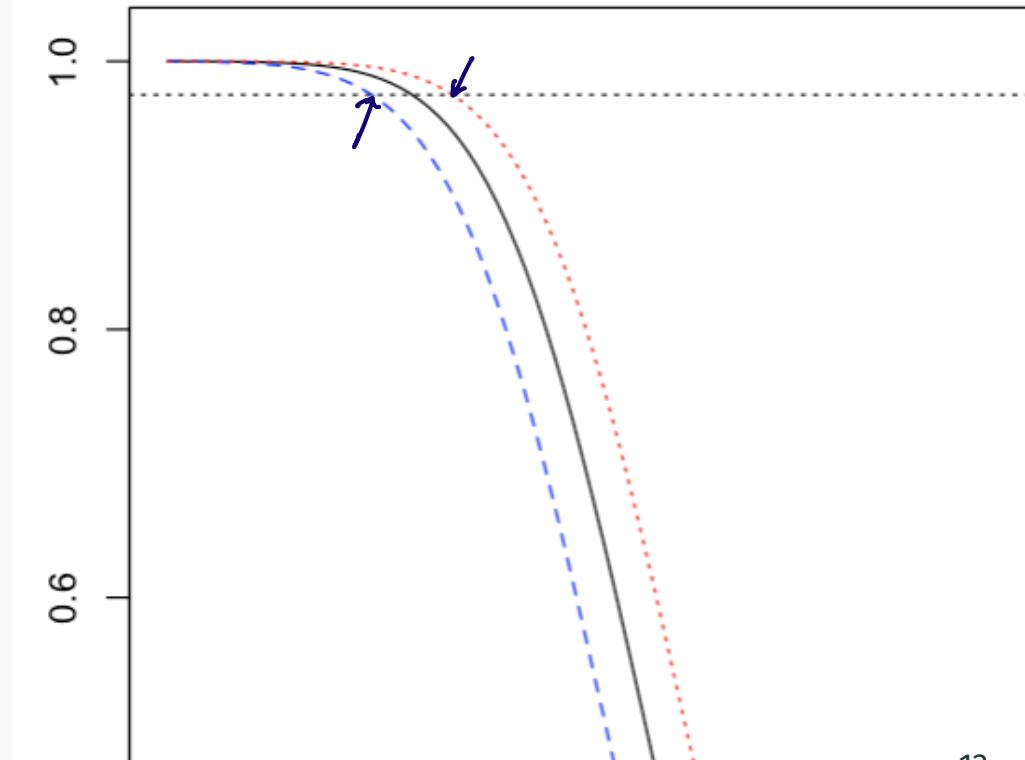
MS 7.2



$$\chi_i \sim N(\theta_i, \frac{1}{n}) \quad i=1, \dots, k$$

$$\psi = \sum_{i=1}^k \theta_i^2$$

$n = 100$ $k = 50$



Approximate normality of posterior

$$\bullet X_1, \dots, X_n \sim f(x^n \mid \theta), \quad \theta \sim \pi(\theta), \quad \pi(\theta \mid x^n) = \frac{f(x^n \mid \theta)}{f(x^n)} \quad x^n = (x_1, \dots, x_n)$$

Approximate normality of posterior

$$\bullet X_1, \dots, X_n \sim f(x^n | \theta), \quad \theta \sim \pi(\theta), \quad \boxed{\pi(\theta | x^n) = \frac{f(x^n | \theta) \pi(\theta)}{f(x^n)}} \quad x^n = (x_1, \dots, x_n)$$

$$\bullet \pi(\theta | x^n) \approx N\{\hat{\theta}, j^{-1}(\hat{\theta})\}; \quad \pi(\theta | x^n) \approx N\{\tilde{\theta}, \tilde{j}(\tilde{\theta})\} \quad \tilde{\theta} = \arg \max_{\theta} \pi(\theta | x^n)$$

\uparrow \uparrow
 prior washed out
 by data

\uparrow
 for large n

$$\tilde{j}(\tilde{\theta}) = -\frac{\partial^2}{\partial \theta^2} \log \pi(\theta | x^n) \quad \theta = \tilde{\theta}$$

$$e^{-\frac{\tilde{j}(\tilde{\theta})}{2} (\theta - \tilde{\theta})^2} \frac{1}{\sqrt{2\pi}} \tilde{j}(\tilde{\theta})^{1/2}$$

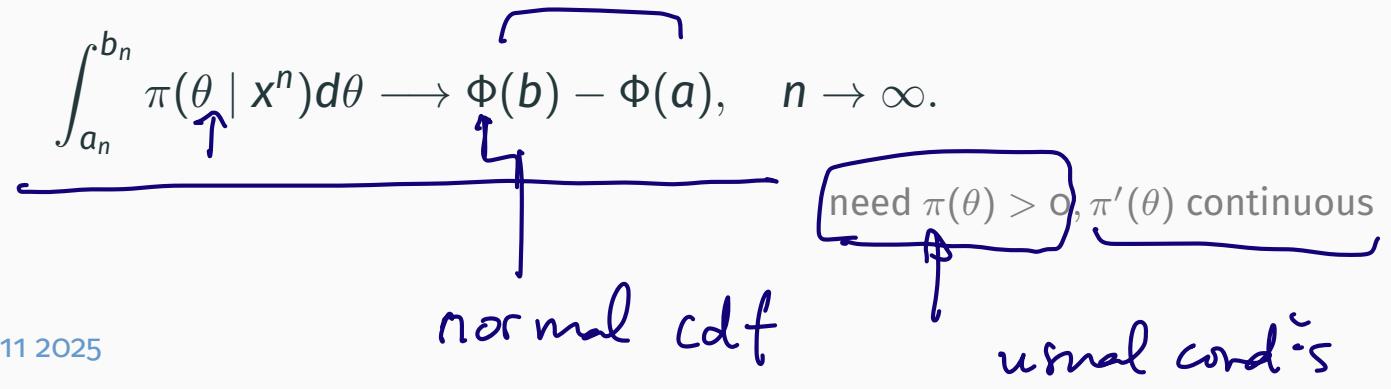
Approximate normality of posterior

- $X_1, \dots, X_n \sim f(x^n | \theta), \quad \theta \sim \pi(\theta), \quad \pi(\theta | x^n) = \frac{f(x^n | \theta)}{f(x^n)} \quad x^n = (x_1, \dots, x_n)$
- $\pi(\theta | x^n) \approx N\{\hat{\theta}, j^{-1}(\hat{\theta})\}; \quad \pi(\theta | x^n) \approx N\{\tilde{\theta}, \tilde{j}(\tilde{\theta})\}$
- careful statement
 - For any $a, b \in \mathbb{R}, a < b$
 - let $a_n = \hat{\theta}_n + aj^{-1/2}(\hat{\theta}_n), b_n = \hat{\theta}_n + bj^{-1/2}(\hat{\theta}_n)$
 - $\hat{\theta}_n$ is the solution of $\ell'(\theta; x^n) = 0$, assumed unique, and $j(\theta) = -\ell''(\theta; x^n)$

$$\underbrace{\hat{\theta} + \hat{a} \text{se}}_{a_n} \leq \theta \leq \underbrace{\hat{\theta}_n + \hat{b} \text{se}}_{b_n}$$

Berger, 1985; Ch.4

Then



Approximate normality of posterior

on $f(x; \theta)$

$$\bullet X_1, \dots, X_n \sim f(x^n | \theta), \quad \theta \sim \pi(\theta), \quad \pi(\theta | x^n) = \frac{f(x^n | \theta)}{f(x^n)} \quad x^n = (x_1, \dots, x_n)$$

$$\bullet \pi(\theta | x^n) \approx N\{\hat{\theta}, j^{-1}(\hat{\theta})\};$$

$$\pi(\theta | x^n) \approx N\{\tilde{\theta}, \tilde{j}(\tilde{\theta})\}$$

used Normal approx.

- approximate posterior probability intervals

$$\hat{\theta} \pm j^{-1}(\hat{\theta}) 1.96 - \text{freq approx 95\% CI}$$

$$\hat{\theta} \pm j^{-1}(\hat{\theta}) 1.96 - \text{approx Bayes credible interval}$$

$$\int_{a_n}^{b_n} \pi(\theta | \underline{x}) d\theta \approx \int_{a_n}^{b_n} e^{\ell(\theta; \underline{x})} \pi(\theta) d\theta \frac{\iint_{a_n}^{b_n} e^{\ell(\theta; \underline{x})} \pi(\theta) d\theta dx}{\int_{a_n}^{b_n} \pi(\theta) d\theta}$$

i.e. $\approx \int_{a_n}^{b_n} e^{\ell(\theta; \underline{x})} \frac{\pi(\theta) d\theta}{\pi(\theta)} = \int_{a_n}^{b_n} e^{\frac{\ell(\theta; \underline{x}) + \log \pi(\theta)}{g(\theta)}} g(\theta) d\theta$

$$= \int_{a_n}^{b_n} e^{\ell(\hat{\theta}; \underline{x}) + (\theta - \hat{\theta}) \ell'(\hat{\theta}; \underline{x}) + \frac{1}{2} (\theta - \hat{\theta})^2 \ell''(\hat{\theta}) + R_n} \pi(\theta) d\theta$$

$$= e^{\ell(\hat{\theta}; \underline{x})} \int_{a_n}^{b_n} e^{-\frac{1}{2} (\theta - \hat{\theta})^2 (-\ell''(\hat{\theta}))} \left\{ \pi(\hat{\theta}) + (\theta - \hat{\theta}) \pi'(\hat{\theta}) + R_2 \right\} d\theta$$

$$\approx \pi(\hat{\theta}) e^{\ell(\hat{\theta}; \underline{x})} \cdot \int_{a_n}^{b_n} e^{-\frac{1}{2} (\theta - \hat{\theta})^2 J(\hat{\theta})} \left\{ 1 + (\theta - \hat{\theta}) \frac{\pi'(\hat{\theta})}{\pi(\hat{\theta})} \right\} d\theta$$

$$\int_{a_n}^{b_n} \pi(\theta | \underline{x}) d\theta = \frac{\sqrt{2\pi}}{|J(\hat{\theta})|^{1/2}} \pi(\hat{\theta}) e^{\ell(\hat{\theta}; \underline{x})} \cdot \int_{a_n}^{b_n} \frac{|J(\hat{\theta})|^{1/2}}{\sqrt{2\pi}} e^{-\frac{1}{2} (\theta - \hat{\theta})^2 J(\hat{\theta})} \left\{ 1 + \dots \right\} d\theta$$

$$z = (\theta - \hat{\theta}) J^{-1/2} (\hat{\theta})$$

$$\text{then } \theta = a_n = \hat{\theta} + a J^{-1/2}$$

$$= \underset{\substack{\uparrow \\ \text{constant in } \theta}}{\text{stuff}} \times \int_a^b \frac{1}{\sqrt{2\pi}} e^{-\frac{1}{2} z^2} dz \left\{ 1 + \text{other stuff} \right\}$$

must be to this
order of approx
constant = 0

for a better version of this

argument \rightarrow Laplace approx =

$$\hat{\pi}(\theta | \underline{x}) = \frac{1}{\sqrt{2\pi}} |J(\hat{\theta})|^{1/2} \cdot e^{\ell(\theta) - \ell(\hat{\theta})} \pi(\hat{\theta})$$

Example: vaccine efficacy

$$\pi(\theta|x) = \pi(\theta|x) \cdot \text{prior}(\theta)$$

Guardian, Jan 24 2021

[Link to Guardian](#)

Pfizer-BioNTech vaccine trial:

vaccine: 22000 subjects, 8 cases

placebo: 22000 subjects, 162 cases

$8/162 = 5\% \Rightarrow 95\%$ efficacy

data released November 18 2020 [link](#)

published December 31 2020 in NEJM [link](#)

Behind the numbers: what does it mean if a Covid vaccine has '90% efficacy'?

David Spiegelhalter and Anthony Masters

Confusion surrounds the vaccines' effectiveness. The leading Cambridge professor clarifies the data behind the trials



▲ People rest in Salisbury Cathedral, England, after receiving the Pfizer/BioNTech vaccine. Photograph: Neil Hall/EPA

Editor's Note: This article was published on December 10, 2020, at NEJM.org.

ORIGINAL ARTICLE

Safety and Efficacy of the BNT162b2 mRNA Covid-19 Vaccine

Fernando P. Polack, M.D., Stephen J. Thomas, M.D., Nicholas Kitchin, M.D., Judith Absalon, M.D., Alejandra Gurtman, M.D., Stephen Lockhart, D.M., John L. Perez, M.D., Gonzalo Pérez Marc, M.D., Edson D. Moreira, M.D., Cristiano Zerbini, M.D., Ruth Bailey, B.Sc., Kena A. Swanson, Ph.D., *et al.*, for the C4591001 Clinical Trial Group*

Article Figures/Media Metrics December 31, 2020
N Engl J Med 2020; 383:2603-2615
DOI: 10.1056/NEJMoa2034577
Chinese Translation 中文翻译

13 References 263 Citing Articles Letters

Results: A total of 43,548 participants underwent randomization, of whom 43,448 received injections: 21,720 with BNT162b2 and 21,728 with placebo. There were 8 cases of Covid-19 with onset at least 7 days after the second dose among participants assigned to receive BNT162b2 and 162 cases among those assigned to placebo; BNT162b2 was 95% effective in preventing Covid-19 (95% credible interval, 90.3 to 97.6).

Table 2. Vaccine Efficacy against Covid-19 at Least 7 days after the Second Dose.*

Efficacy End Point	BNT162b2		Placebo		Vaccine Efficacy, % (95% Credible Interval)‡	Posterior Probability (Vaccine Efficacy >30%)§
	No. of Cases	Surveillance Time (n)†	No. of Cases	Surveillance Time (n)†		
		(N=18,198)		(N=18,325)		
Covid-19 occurrence at least 7 days after the second dose in participants without evidence of infection	8	2.214 (17,411)	162	2.222 (17,511)	95.0 (90.3–97.6)	>0.9999
		(N=19,965)		(N=20,172)		
Covid-19 occurrence at least 7 days after the second dose in participants with and those without evidence of infection	9	2.332 (18,559)	169	2.345 (18,708)	94.6 (89.9–97.3)	>0.9999

* The total population without baseline infection was 36,523; total population including those with and those without prior evidence of infection was 40,137.

† The surveillance time is the total time in 1000 person-years for the given end point across all participants within each group at risk for the end point. The time period for Covid-19 case accrual is from 7 days after the second dose to the end of the surveillance period.

‡ The credible interval for vaccine efficacy was calculated with the use of a beta-binomial model with prior beta (0.700102, 1) adjusted for the surveillance time.

§ Posterior probability was calculated with the use of a beta-binomial model with prior beta (0.700102, 1) adjusted for the surveillance time.

Credible intervals

- vaccine group 18000 participants; 8 cases
- placebo group 18000 participants; 162 cases
- $0.05 = 8/162 \rightarrow 95\% \text{ efficacy}$
- model
 $X_1 \sim \underline{\text{Poisson}}(\lambda\psi), \quad X_2 \sim \underline{\text{Poisson}}(\lambda) \quad \underline{X_1 | S = X_1 + X_2 \sim \text{Binom}(S, \psi/(1 + \psi))}$
- prior $\text{Beta}(a, b) \rightarrow \text{posterior } \text{Beta}(x_1 + a, s - x_1 + b)$

Credible intervals

- vaccine group 18000 participants; 8 cases
- placebo group 18000 participants; 162 cases
- $0.05 = 8/162 \rightarrow 95\% \text{ efficacy}$

- model

$$X_1 \sim \text{Poisson}(\lambda\psi), \quad X_2 \sim \text{Poisson}(\lambda)$$

- prior $\text{Beta}(a, b) \rightarrow \text{posterior } \text{Beta}(x_1 + a, s - x_1 + b)$

$$8 + 0.7 \quad 162 + 1$$

- qbeta(c(0.025, 0.975), shape1 = 8.7, shape2 = 163)

- > [1] 0.02319 0.08799

- $1 - \text{Last.value}/(1 - \text{Last.value})$

- > [1] 0.97625 0.90352

credible interval

$$X_1 | S = X_1 + X_2 \sim \text{Binom}(S, \psi/(1 + \psi))$$

170 : u

$$VE = 1 - \psi$$

- multiparameter model $f(\mathbf{X}; \boldsymbol{\theta})$

$$\text{pr}_{\boldsymbol{\theta}}\{\boldsymbol{\theta} \in R(\mathbf{X})\} \geq 1 - \alpha,$$

for all $\boldsymbol{\theta}$, with equality for some $\boldsymbol{\theta}$

- pivotal method:

$$1 - \alpha = \text{pr}_{\boldsymbol{\theta}}\{a \leq g(\mathbf{X}; \boldsymbol{\theta}) \leq b\} = \text{pr}_{\boldsymbol{\theta}}\{\boldsymbol{\theta} \in R(\mathbf{X})\}$$

- Example: $\underline{\mathbf{X}_1, \dots, \mathbf{X}_n}$ i.i.d. $N_p(\mu, \Sigma)$

$$\boldsymbol{\theta} \in \mathbb{R}^P$$

MS Ex.7.8

- exact pivot

$$g(\mathbf{X}; \underline{\mu}) = \frac{n(n-p)}{p(n-1)} (\underline{\hat{\mu}} - \underline{\mu})^T \underline{\hat{\Sigma}}^{-1} (\underline{\hat{\mu}} - \underline{\mu})$$

"Student's T"

$$\hat{\Sigma} = \frac{1}{n-1} \sum_{i=1}^n (\underline{x}_i - \underline{\bar{x}})(\underline{x}_i - \underline{\bar{x}})^T$$

$\sim F_{p, n-p}$ d.f.

 Hotelling's T²

- multiparameter model $f(\mathbf{X}; \theta)$

$$\text{pr}_\theta\{\boldsymbol{\theta} \in R(\mathbf{X})\} \geq 1 - \alpha,$$

for all θ , with equality for some θ

- pivotal method:

$$1 - \alpha = \text{pr}_\theta\{a \leq g(\mathbf{X}; \theta) \leq b\} = \text{pr}_\theta\{\boldsymbol{\theta} \in R(\mathbf{X})\}$$

- Example: $\mathbf{X}_1, \dots, \mathbf{X}_n$ i.i.d. $N_p(\mu, \Sigma)$
- exact pivot

MS Ex.7.8

$$g(\mathbf{X}; \mu) = \frac{n(n-p)}{p(n-1)} (\hat{\mu} - \mu)^T \hat{\Sigma}^{-1} (\hat{\mu} - \mu)$$

conf. region
exact

$$R(\mathbf{X}) = \{\mu : \underbrace{\frac{n(n-p)}{p(n-1)} (\hat{\mu} - \mu)^T \hat{\Sigma}^{-1} (\hat{\mu} - \mu)}_{\equiv} \leq f_{1-\alpha}\}$$

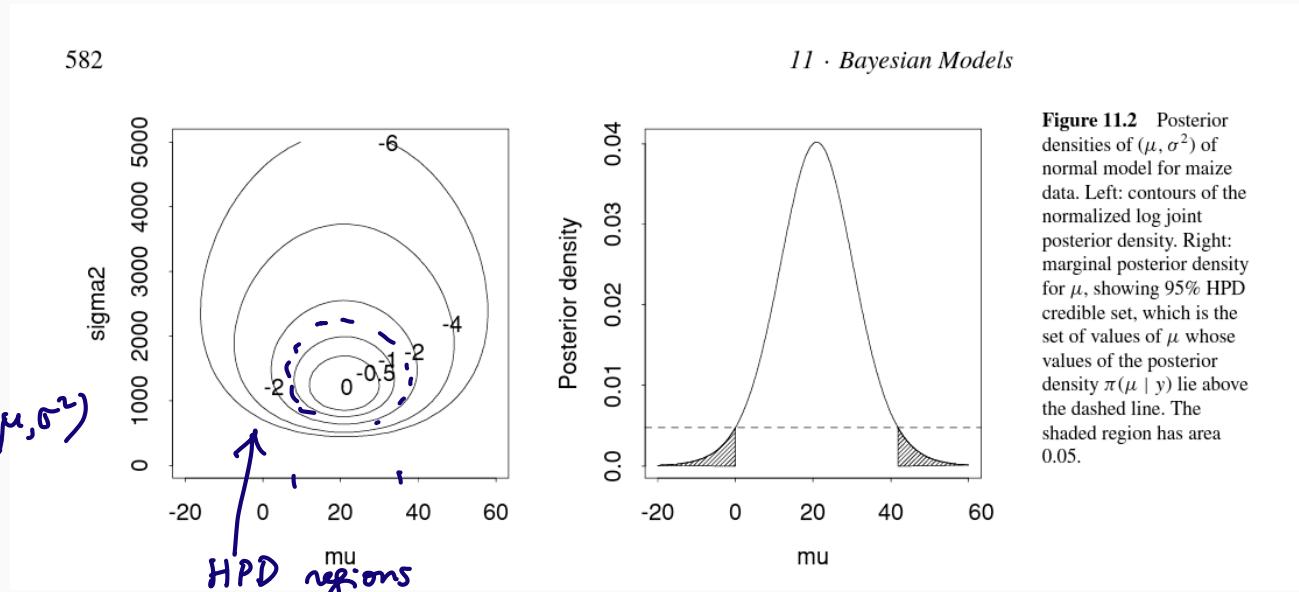
 $F_{p,n-p}$

Highest posterior density (HPD) regions

SM 11.2.1

- HPD region C for θ :

$$(1) \quad \int_C \pi(\theta | \mathbf{x}) = 1 - \alpha$$
$$(2) \quad \pi(\theta | \mathbf{x}) \geq \pi(\theta^* | \mathbf{x})$$



Approximate confidence regions

- maximum likelihood estimator is approximately normal

-

$$\widehat{\theta} \sim N\{\theta, I_n^{-1}(\widehat{\theta})\} \implies (\widehat{\theta} - \theta)^T I_n(\widehat{\theta})(\widehat{\theta} - \theta) \sim \chi_k^2$$

-

$$1 - \alpha \approx \text{pr}_{\theta}\{\theta \in R(\widehat{\theta})\}$$

-

$$R(\widehat{\theta}) = \{\theta : (\widehat{\theta} - \theta)^T I_n(\widehat{\theta})(\widehat{\theta} - \theta) \leq \chi_{k,1-\alpha}^2\}$$

$$\theta \in \mathbb{R}^p$$

Approximate confidence regions

- maximum likelihood estimator is approximately normal

-

$$\hat{\theta} \sim N\{\theta, I_n^{-1}(\hat{\theta})\} \implies (\hat{\theta} - \theta)^T I_n(\hat{\theta})(\hat{\theta} - \theta) \sim \chi_k^2$$

-

$$1 - \alpha \approx \text{pr}_{\theta}\{\theta \in R(\hat{\theta})\}$$

-

$$R(\hat{\theta}) = \{\theta : (\hat{\theta} - \theta)^T I_n(\hat{\theta})(\hat{\theta} - \theta) \leq \chi_{k,1-\alpha}^2\}$$

- $k = 1$:

$$\hat{\theta} \pm z_{1-\alpha/2} \widehat{se}(\hat{\theta})$$

AoS Thm 6.16

Likelihood ratio based approximate confidence regions

- $X_1, \dots, X_n \sim f(\mathbf{x}; \theta)$
- $L(\theta; \mathbf{x}) = f(\mathbf{x}; \theta), \quad \ell(\theta) = \log L(\theta; \mathbf{x})$
- $w(\theta) = 2\{\ell(\hat{\theta}) - \ell(\theta)\} \xrightarrow{d} \chi_p^2, \quad n \rightarrow \infty \quad \leftarrow \text{approx probab}$

freq. v. of
HPD
interval

$$\Pr\{w(\theta) \leq \chi_p^2(1-\alpha)\} = 1-\alpha$$

defines conf. region for $\theta \in \mathbb{R}^P$

Likelihood ratio based approximate confidence regions

- $X_1, \dots, X_n \sim f(\mathbf{x}; \theta)$
- $L(\theta; \mathbf{x}) = f(\mathbf{x}; \theta), \quad \ell(\theta) = \log L(\theta; \mathbf{x})$
- $w(\theta) = 2\{\ell(\hat{\theta}) - \ell(\theta)\} \xrightarrow{d} \chi_p^2, \quad n \rightarrow \infty$

- approximation:

$$w(\theta) \stackrel{\sim}{\sim} \chi_p^2$$

- approximate confidence region

$$\{\theta : w(\theta) \leq \chi_{p,1-\alpha}^2\}$$

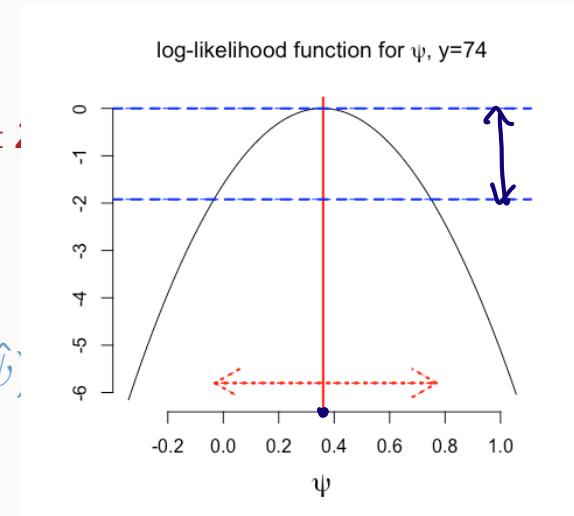
- model $Y \sim f(y; \psi, \lambda)$, $\psi \in \mathbb{R}, \lambda \in \mathbb{R}^{d-1}$, $\theta = (\psi, \lambda)$ $y = (y_1, \dots, y_n)$
- log-likelihood function $\ell(\psi, \lambda; y) = \log f(y; \psi, \lambda) = \sum \log f(y_i; \psi, \lambda)$ if independent
- profile log-likelihood function $\ell_p(\psi) = \ell(\psi, \hat{\lambda}_\psi)$ maximize over λ
- maximum likelihood estimate $j_p(\psi) = -\ell''_p(\psi)$
 $\hat{\psi} \stackrel{d}{\sim} N\{\psi, j_p^{-1/2}(\psi)\} \implies 1 - \alpha \text{ CI} \approx \hat{\psi} \pm z_{1-\alpha/2} \hat{j}_p^{1/2}$
- likelihood ratio test
 $2\{\ell_p(\hat{\psi}) - \ell_p(\psi)\} \stackrel{d}{\sim} \chi^2_1 \implies 1 - \alpha \text{ CI} \approx \{\psi : 2\{\ell_p(\hat{\psi}) - \ell_p(\psi)\} \leq \chi^2_{1,1-\alpha}\}$

- model $Y \sim f(y; \psi, \lambda)$, $\psi \in \mathbb{R}, \lambda \in \mathbb{R}^{d-1}$, $\theta = (\psi, \lambda)$ $y = (y_1, \dots, y_n)$
- log-likelihood function $\ell(\psi, \lambda; y) = \log f(y; \psi, \lambda) = \sum \log f(y_i; \psi, \lambda)$ if independent
- profile log-likelihood function $\ell_p(\psi) = \ell(\psi, \hat{\lambda}_\psi)$ maximize over λ
- maximum likelihood estimate

$$\hat{\psi} \stackrel{\text{d}}{\sim} N\{\psi, j_p^{-1/2}(\psi)\} \implies 1 - \alpha \text{ CI} \approx \hat{\psi} \pm z_{\alpha/2} j_p^{-1/2}(\hat{\psi})$$

- likelihood ratio test

$$2\{\ell_p(\hat{\psi}) - \ell_p(\psi)\} \stackrel{\text{d}}{\sim} \chi^2_1 \implies 1 - \alpha \text{ CI} \approx \{\psi : 2\{\ell_p(\hat{\psi}) - \ell_p(\psi)\} \leq \chi^2_{1-\alpha}\}$$



Nonparametric confidence bands

AoS 7.4

- recall X_1, \dots, X_n , i.i.d. $F(\cdot)$
- empirical cdf

nonp. *unk.*

est.

$$\hat{F}_n(t) = \frac{1}{n} \sum_{i=1}^n \mathbf{1}\{X_{(i)} \leq t\}$$

- properties:

$$E\{\hat{F}_n(t)\} = F(t), \quad \text{var}\{\hat{F}_n(t)\} = \frac{1}{n} F(t)\{1 - F(t)\}$$

- pointwise approximate confidence limits $\hat{F}_n(t) \pm z_{1-\alpha/2} [\hat{F}_n(t)\{1 - \hat{F}_n(t)\}]^{1/2}$

any fixed t

binomial approx. prob

- recall $X_1, \dots, X_n, i.i.d. F(\cdot)$

- empirical cdf

$$\hat{F}_n(t) = \frac{1}{n} \sum_{i=1}^n \mathbf{1}\{X_{(i)} \leq t\}$$

- properties:

Bonferroni $E\{\hat{F}_n(t)\} = F(t), \quad \text{var}\{\hat{F}_n(t)\} = \frac{1}{n}F(t)\{1 - F(t)\}$

any fixed t

- pointwise approximate confidence limits $\hat{F}_n(t) \pm z_{1-\alpha/2}[\hat{F}_n(t)\{1 - \hat{F}_n(t)\}]^{1/2}$

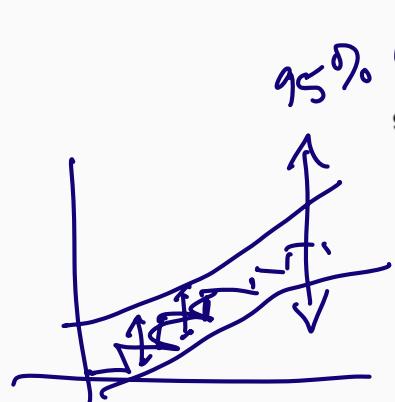
- simultaneous confidence band: $\text{pr}\{\underline{L}(t) \leq F(t) \leq \underline{U}(t) \text{ for all } t\} \geq 1 - \alpha$:

$$\underline{L}(t) = \max\{\hat{F}_n(t) - \epsilon_n, 0\}, \quad \underline{U}(t) = \min\{\hat{F}_n(t) + \epsilon_n, 1\}, \quad \epsilon_n = \left\{ \frac{1}{2n} \log \left(\frac{2}{\alpha} \right) \right\}^{1/2}$$

best L, U

... Nonparametric confidence bands

AoS 7.4



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7. Estimating the CDF and Statistical Functionals

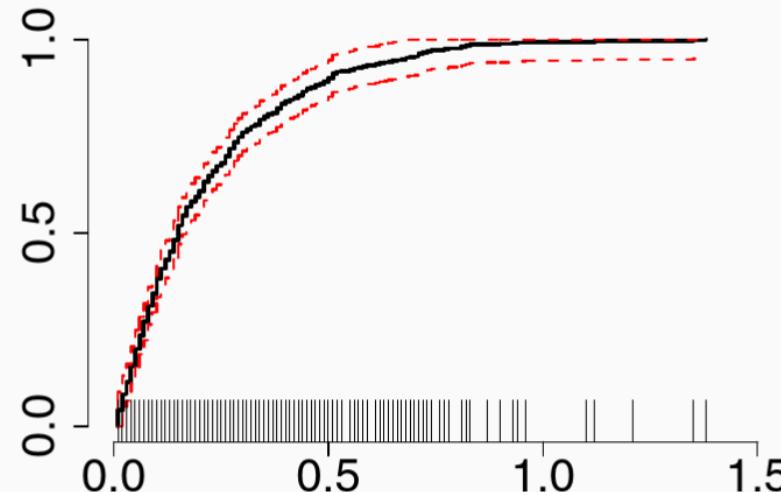
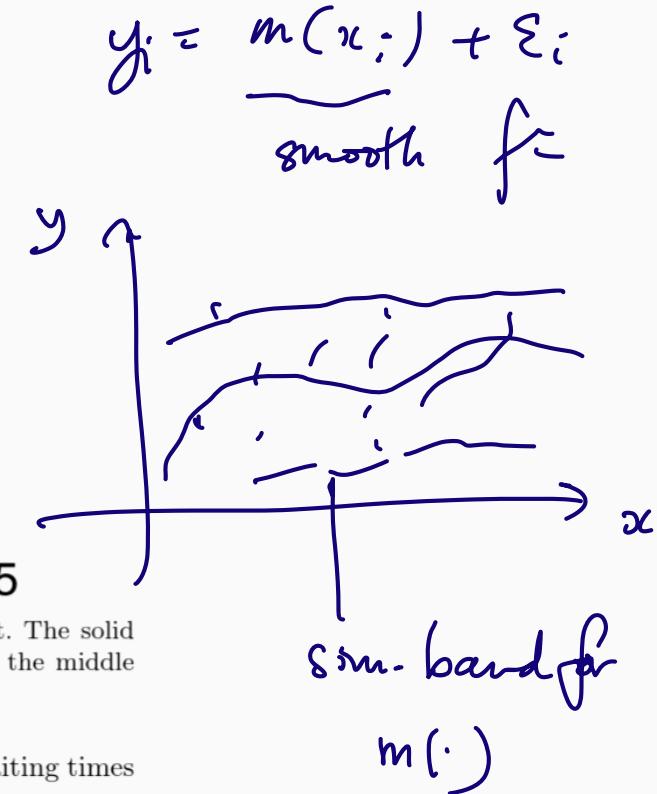


FIGURE 7.1. Nerve data. Each vertical line represents one data point. The solid line is the empirical distribution function. The lines above and below the middle line are a 95 percent confidence band.



7.2 Example (Nerve Data). Cox and Lewis (1966) reported 799 waiting times between successive pulses along a nerve fiber. Figure 7.1 shows the empirical CDF \hat{F}_n . The data points are shown as small vertical lines at the bottom of the plot. Suppose we want to estimate the fraction of waiting times between .4 and .6 seconds. The estimate is $\hat{F}_n(.6) - \hat{F}_n(.4) = .93 - .84 = .09$. ■

Example: Bootstrap confidence intervals

```
> alpha = 0.05  
>  
> # Normal-based CI  
> c(phat - qnorm(1-alpha/2)*sd(bs_est),  
+    phat + qnorm(1-alpha/2)*sd(bs_est))  
[1] 0.05709686 0.44290314  
>  
> # Percentile CI  
> c(quantile(bs_est, alpha/2),  
+    quantile(bs_est, 1-alpha/2))  
2.5% 97.5%  
0.10 0.45
```

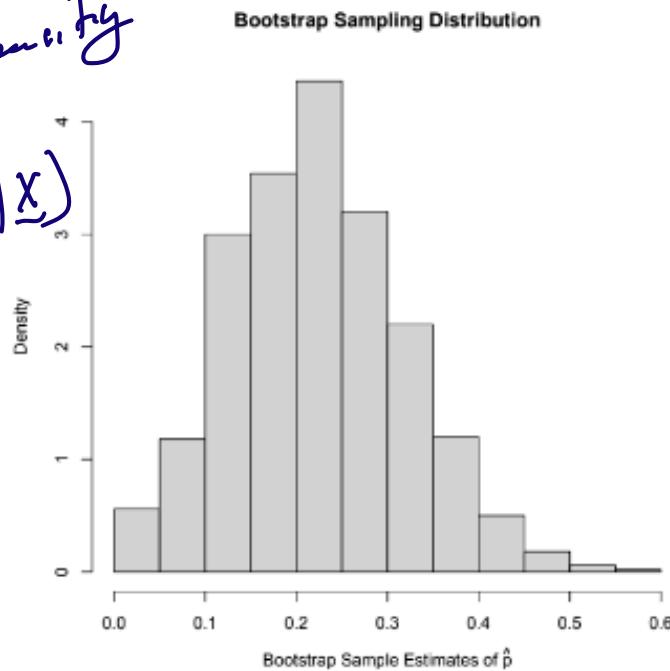
Calculate in R

$$\Rightarrow n-z_0 = \begin{cases} 5 \text{ successes} \\ 15 \text{ failures} \end{cases}$$

Example: Bootstrap confidence intervals

B.S. density

$$p^*(\hat{\psi}^* | \bar{x})$$



$$\tilde{X}^* \sim i.i.d f(x_i; \hat{\theta})$$

$$\hat{F}_n(\cdot)$$

$$\psi = g(\theta)$$

