Contents lists available at ScienceDirect

Journal of Multivariate Analysis

journal homepage: www.elsevier.com/locate/jmva

Uniformly valid inference based on the Lasso in linear mixed models

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ARTICLE INFO

Article history: Received 27 May 2022 Received in revised form 1 August 2023 Accepted 2 August 2023 Available online 9 August 2023

AMS 2020 subject classifications: primary 62F25 secondary 62J10 62J07

Keywords: Confidence sets REML Sparsity Variance components Uniform consistency

ABSTRACT

Linear mixed models (LMMs) are suitable for clustered data and are common in biometrics, medicine, survey statistics, and many other fields. In those applications, it is essential to carry out valid inference after selecting a subset of the available variables. We construct confidence sets for the fixed effects in Gaussian LMMs that are based on Lasso-type estimators. Aside from providing confidence regions, this also allows for quantification of the joint uncertainty of both variable selection and parameter estimation in the procedure. To show that the resulting confidence sets for the fixed effects are uniformly valid over the parameter spaces of both the regression coefficients and the covariance parameters, we also prove the novel result on uniform Cramér consistency of the restricted maximum likelihood (REML) estimators of the covariance parameters. The superiority of the constructed confidence sets to naïve postselection procedures is validated in simulations and illustrated with a study of the acid-neutralization capacity of lakes in the United States.

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1. Introduction

Linear mixed models (LMMs) are regression models that incorporate dependency structures that occur in clustered data. They are widely applied in many empirical sciences ranging from genetics [11] to survey statistics [28] and many more. Comprehensive reviews can be found in [8,29]. The standard LMM can be written as

$$\mathbf{y}_{i} = \mathbf{X}_{i}\boldsymbol{\beta}_{0} + \mathbf{Z}_{i}\mathbf{v}_{i} + \boldsymbol{\varepsilon}_{i}, \quad i \in \{1, \dots, m\}, \quad \boldsymbol{\varepsilon}_{i} \sim \mathcal{N}_{n_{i}}\{\mathbf{0}_{n_{i}}, \boldsymbol{\Omega}_{i}(\boldsymbol{\theta}_{0})\}, \quad \mathbf{v}_{i} \sim \mathcal{N}_{q}\{\mathbf{0}_{q}, \boldsymbol{\Psi}(\boldsymbol{\theta}_{0})\}, \quad (1)$$

with observations $\mathbf{y}_i \in \mathbb{R}^{n_i}$, known fixed covariates $\mathbf{X}_i \in \mathbb{R}^{n_i \times p}$, $\mathbf{Z}_i \in \mathbb{R}^{n_i \times q}$, and $\mathbf{v}_i \in \mathbb{R}^q$ and $\mathbf{\varepsilon}_i \in \mathbb{R}^{n_i}$ being independent random variables and each independently distributed for all clusters $i \in \{1, ..., m\}$. The regression coefficient vector $\boldsymbol{\beta}_0$ is referred to as fixed effects, whereas \mathbf{v}_i are the random effects. While all observations share the same regression coefficients in the fixed effects, only n_i observations in each cluster share the same realization of \mathbf{v}_i in the random effects. The latter can be associated, e.g., with subject-specific effects in medical trials or group effects in longitudinal studies. The regression coefficients $\boldsymbol{\beta}_0 \in \mathbb{R}^p$ and covariance parameters $\boldsymbol{\theta}_0 \in \boldsymbol{\Theta} \subset \mathbb{R}^r_{>0}$ are unknown and have to be estimated. Consistency of parameter estimators for $\boldsymbol{\beta}_0$ and $\boldsymbol{\theta}_0$ is ensured if the number of clusters *m* tends to infinity, whereas the sample size per cluster n_i may stay bounded or grow as well [8, Section 3.6.2].

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https://doi.org/10.1016/j.jmva.2023.105230





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In practice, often both model selection and estimation need to be performed. Naïve two-stage methods for inference in regression models consist of a variable selection step, which is based, e.g., on an information criterion, and subsequent estimation of the parameters and inference in the obtained model in a second step. However, this naïve inferential theory ignores the first model selection step and does not account for the additional uncertainty introduced by the selection procedure. This issue has been laid out in detail in [30]. In recent years, different approaches to this problem have emerged in the literature. The PoSi or post-selection inference framework of [1] provides a general procedure to construct confidence regions for a best-approximation parameter that depends on the selected model. In that regard, a similar perspective was taken by [18] who also aim to cover this surrogate quantity rather than a 'true' parameter and introduce the idea of selective inference with a focus on the Lasso. This selective inference approach, where the coverage holds conditionally on the selected model, is followed in [34] specifically for LMMs. Following a similar paradigm in the sense of conditional inference, [4] suggest an approach based on the Akaike Information Criterion (AIC) which is carried out for LMMs in [5]. Contrasting this, a more classical perspective is taken in the general debiasing approach of [39] and related articles, e.g., [21] for LMMs, where the target is (a sub-vector of) the true parameter and the coverage probability holds unconditionally. We follow this classical approach in that we also consider unconditional inference with the aim to cover the true parameter. All the methods listed above are motivated from a high-dimensional perspective. In our work, we focus on a low-dimensional setting, that is, when the number of variables is smaller than the number of observations.

In this work, we employ the least absolute shrinkage and selection operator (Lasso) [37], which offers a single step approach to simultaneous variable selection and estimation for the regression coefficients. Although in the context of the linear mixed models, the Lasso can be applied to the selection of both fixed and random effects [2,13,25,27], we consider selection and inference for β_0 only. Selection of covariates β_0 is fundamental to interpreting the association between the set of covariates and the response, which is of high relevance in most practical problems.

The usefulness and wide application of Lasso sparked interest in how to construct confidence intervals based on the resulting Lasso estimators. Not too surprisingly, the model selection step inherent to the Lasso estimator leads to challenges with respect to inference. For the Lasso and related estimators, this manifests in the general difficulty that its finite sample and, in an appropriate framework, also the asymptotic distribution depends on the unknown coefficient parameters β_0 in an intricate manner [32]. This is problematic since honest confidence sets in the sense of [20] have to attain nominal coverage over the whole parameter space, which has been investigated in detail for the Lasso in the case of orthogonal covariates in [33]. For a general low-dimensional framework, where the number of parameters is smaller than the sample size, [9] have obtained exact and uniformly valid confidence sets. All these references consider ordinary linear models and do not cover the case of linear mixed models.

The contribution of the present article to the LMM framework is two-fold. First, we show that the covariance parameter θ_0 can be estimated Cramér-consistently in a uniform manner. Second, we construct confidence regions for β_0 based on the Lasso which are uniformly valid over the space of coefficient and covariance parameters. Unlike an ordinary linear model, an LMM requires the estimation of covariance parameters θ_0 in addition to the coefficient parameters β_0 . The key idea is to estimate the former with restricted maximum likelihood (REML) as this procedure allows to separate the estimation of θ_0 and β_0 . A novel uniform consistency result for the REML estimators is established, extending the previous result of [41]. This new result is crucial to non-trivially extend the reasoning of [9] to construct confidence sets for β_0 that hold the coverage not only uniformly over the space of coefficient parameters but also over the space of covariance parameters. Thus, researchers are provided with a method that allows valid inference based on the Lasso in LMMs. Our analysis is provided within a low-dimensional framework where p < n.

The remainder of the article is structured as follows. First, we specify the setting and regularity conditions and state the estimation procedure for the fixed effects β_0 in Section 2. Next, in Section 3, the estimation of covariance parameters θ_0 is discussed and its uniform consistency is established. In Section 4, the confidence regions for β_0 based on the Lasso are presented. Their usefulness and limitations, and, in particular, their superiority to naïve approaches is demonstrated in a simulation study in Section 5. A real data example is provided in Section 6. The article concludes with a discussion in Section 7.

Throughout the remainder of the article, we use the following notation. For a matrix **A**, tr(**A**) and rk(**A**) denote the trace and rank of **A**, respectively. By $\eta_i(\mathbf{A})$, we refer to the *i*th eigenvalue of **A**, sorted in descending order. The symbol $\|\mathbf{A}\|$ denotes the Frobenius norm, i.e., $\|\mathbf{A}\|^2 = \text{tr}(\mathbf{A}^\top \mathbf{A})$. Finally, for $\mathbf{A}_i \in \mathbb{R}^{n_i \times m_i}$, blockdiag(\mathbf{A}_i)_{i=1}^n denotes the $(\sum_{i=1}^n n_i \times \sum_{i=1}^n m_i)$ -dimensional block-diagonal matrix with \mathbf{A}_i along the diagonal block and zeros elsewhere.

2. Setting and regularity conditions

Consider a linear model with $n \in \mathbb{N}$ dependent observations and model equation

$$\mathbf{y} = \mathbf{X}\boldsymbol{\beta}_0 + \boldsymbol{\epsilon}, \quad \boldsymbol{\epsilon} \sim \mathcal{N}_n \{\mathbf{0}_n, \mathbf{V}(\boldsymbol{\theta}_0)\}.$$

(2)

The covariance matrix $\mathbf{V}(\cdot) \in \mathbb{R}^{n \times n}$ models the dependency amongst the observations. The vectors $\boldsymbol{\beta}_0 \in \mathbb{R}^p$ as well as $\boldsymbol{\theta}_0 \in \boldsymbol{\Theta} \subset \mathbb{R}^r_{>0}$ are unknown and remain to be estimated. Model (1) can be represented as (2) with $n = \sum_{i=1}^m n_i$,

 $\mathbf{y} = (\mathbf{y}_{n_1}, \dots, \mathbf{y}_{n_m})^{\top}, \mathbf{X} = (\mathbf{X}_1^{\top}, \dots, \mathbf{X}_m^{\top})^{\top}$ and $\mathbf{V}(\boldsymbol{\theta}_0) = \text{blockdiag}\{\mathbf{Z}_i \boldsymbol{\Psi}(\boldsymbol{\theta}_0) \mathbf{Z}_i^{\top} + \boldsymbol{\Omega}_i(\boldsymbol{\theta}_0)\}_{i=1}^m$. For given tuning parameters $\lambda_1, \ldots, \lambda_n$ and a REML estimator $\hat{\theta}$ for θ_0 (defined by (5) in Section 3), consider the Lasso estimator

$$\hat{\boldsymbol{\beta}}_{L} = \underset{\boldsymbol{\beta} \in \mathbb{R}^{p}}{\operatorname{argmin}} \left\{ \left\| \mathbf{V}(\hat{\boldsymbol{\theta}})^{-1/2} \left(\mathbf{y} - \mathbf{X} \boldsymbol{\beta} \right) \right\|^{2} + 2 \sum_{j=1}^{p} \lambda_{j} \left| \beta_{j} \right| \right\}$$
(3)

for β_0 . If $\hat{\theta}$ in (3) were replaced by the true parameter θ_0 , the objective function would reduce to the standard ℓ_1 -penalization of the Lasso. Note that $\hat{\beta}_l$ simultaneously performs estimation and variable selection.

This paper aims to provide confidence regions based on $\hat{\beta}_{L}$ for the fixed effects β_{0} , which hold their coverage probability uniformly over the space of coefficient and covariance parameters. Formally, we find a set $M \subset \mathbb{R}^p$ such that $\inf_{\beta_0,\theta_0} \Pr_{\beta_0,\theta_0} \{\sqrt{n}(\hat{\beta}_L - \beta_0) \in M\} \approx 1 - \alpha$ for some nominal level $\alpha \in (0, 1)$, which is attained uniformly over $\beta_0 \in \mathbb{R}^p$ and $\theta_0 \in \Theta$. We borrow and adapt the notation of [9]. The key idea of that article is the following: Instead of treating $\hat{\beta}_{i}$ with its intractable distribution directly, one considers limiting versions of the objective function in (3), in which each component of β_0 tends to infinity in absolute value. This process gives 2^p minimizers. One can choose M such that it contains each of these minimizers with a probability of at least $1 - \alpha$. It can be shown that this procedure does yield exact and uniformly valid confidence sets for ordinary linear models. For the model in (2) and optimization problem in (3), this gives the 2^p minimizers

$$\hat{\mathbf{u}}_{\mathbf{d}} = \underset{\mathbf{u} \in \mathbb{R}^{p}}{\operatorname{argmin}} \mathbf{u}^{\top} \hat{\mathbf{C}} \mathbf{u} - 2\mathbf{u}^{\top} \hat{\mathbf{w}} + 2\mathbf{u}^{\top} \boldsymbol{\Lambda} \mathbf{d} = \hat{\mathbf{C}}^{-1} \Big(\hat{\mathbf{w}} - \boldsymbol{\Lambda} \mathbf{d} \Big), \tag{4}$$

where $\hat{\mathbf{C}} = n^{-1} \mathbf{X}^{\top} \mathbf{V}(\hat{\boldsymbol{\theta}})^{-1} \mathbf{X}$, $\hat{\mathbf{w}} = n^{-1/2} \mathbf{X}^{\top} \mathbf{V}(\hat{\boldsymbol{\theta}})^{-1} (\mathbf{y} - \mathbf{X} \boldsymbol{\beta}_0)$, $\boldsymbol{\Lambda} = n^{-1/2} \operatorname{diag}(\lambda_1, \dots, \lambda_p)$ and $\mathbf{d} \in \{-1, 1\}^p$. The latter plays the role of representing the signs of the coefficients. The contribution of the current paper is to prove that a similar idea can be used for LMMs, for which the main challenge is to show that the uniform coverage carries over to the covariance parameters and that the estimation of these parameters does not interfere with the required uniformity over the coefficient parameters.

Subsequently, we will denote by \mathbf{u}_d , \mathbf{C} and \mathbf{w} the counterparts of $\hat{\mathbf{u}}_d$, $\hat{\mathbf{C}}$ and $\hat{\mathbf{w}}$, with $\hat{\theta}$ replaced by θ_0 . As the distribution of $\hat{\mathbf{u}}_d$ is not analytically available, the proofs use that $\mathbf{u}_d \sim \mathcal{N}_p(-\mathbf{C}^{-1}\mathbf{\Lambda}\mathbf{d}, \mathbf{C}^{-1})$. We only consider confidence sets of ellipsoidal shape, that is, M will be set to $E(\hat{\mathbf{C}}, t) = \{\mathbf{z} \in \mathbb{R}^p \mid \mathbf{z}^\top \hat{\mathbf{C}} \mathbf{z} \leq t\}, t > 0$. Their specific form is used in Lemma 3 to establish the minimal coverage probability over θ_0 .

The following regularity conditions on the model (2) are imposed.

- (A) For a constant $c \in (1, \infty)$, $\theta_0 \in \Theta = \{ \theta \in \mathbb{R}^r_{>0} \mid \max(\theta) / \min(\theta) \le c \}$.
- (B) $\mathbf{V}(\boldsymbol{\theta}_0) = \sum_{k=1}^r \theta_{0,k} \mathbf{H}_k$ with $\mathbf{H}_k, k \in \{1, \dots, r-1\}$, positive semi-definite and \mathbf{H}_r positive definite. Further, m = 1 $\min_{k\in\{1,\ldots,r\}} \operatorname{rk}(\mathbf{H}_k) \to \infty.$
- (C) $\operatorname{rk}(\mathbf{X}) = p < n$ and for all $\boldsymbol{\theta}_0 \in \Theta$: $\mathbf{C} = n^{-1}\mathbf{X}^\top \mathbf{V}(\boldsymbol{\theta}_0)^{-1}\mathbf{X} \to \tilde{\mathbf{C}}$ pointwise for finite and positive definite $\tilde{\mathbf{C}}$ as $n \to \infty$. (D) For constants $0 < \underline{\omega} \le \overline{\omega} < \infty$ it holds that $\underline{\omega} \le \operatorname{tr}\{(\sum_{l=1}^r \mathbf{H}_l)^{-1}\mathbf{H}_k\}/\operatorname{rk}(\mathbf{H}_k) \le \overline{\omega}$ and $\sum_{j=1}^p \eta_j\{(\sum_{l=1}^r \mathbf{H}_l)^{-1}\mathbf{H}_k\} = O(1)$, $k \in \{1, ..., r\}.$
- (E) For $\mathbf{P}(\boldsymbol{\theta}, a) = \mathbf{V}(\boldsymbol{\theta})^{-1} a\mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{X} \{\mathbf{X}^{\top}\mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{X}\}^{-1}\mathbf{X}^{\top}\mathbf{V}(\boldsymbol{\theta})^{-1} \in \mathbb{R}^{n \times n}, a \geq 1 \text{ and } \mathbf{K} \in \mathbb{R}^{r \times r} \text{ with entries } (\mathbf{K})_{ij} = \mathbf{V}(\boldsymbol{\theta})^{-1} \mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{X} \{\mathbf{X}^{\top}\mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{X}\}^{-1}\mathbf{X}^{\top}\mathbf{V}(\boldsymbol{\theta})^{-1} \in \mathbb{R}^{n \times n}, a \geq 1 \text{ and } \mathbf{K} \in \mathbb{R}^{r \times r} \text{ with entries } (\mathbf{K})_{ij} = \mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{X} \{\mathbf{X}^{\top}\mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{X}\}^{-1}\mathbf{X}^{\top}\mathbf{V}(\boldsymbol{\theta})^{-1} \in \mathbb{R}^{n \times n}, a \geq 1 \text{ and } \mathbf{K} \in \mathbb{R}^{n \times r} \text{ with entries } (\mathbf{K})_{ij} = \mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{X} \{\mathbf{X}^{\top}\mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{X}\}^{-1}\mathbf{X}^{\top}\mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{X} \{\mathbf{X}^{\top}\mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{X}\}^{-1}\mathbf{X}^{\top}\mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{X} \{\mathbf{X}^{\top}\mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{X}\}^{-1}\mathbf{X}^{\top}\mathbf{X}^{\top}\mathbf{X}^{\top}\mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{X}\}^{-1}\mathbf{X}^{\top}\mathbf{X}^{\top}\mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{X} \{\mathbf{X}^{\top}\mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{X}\}^{-1}\mathbf{X}^{\top$ tr{**P**(**1**_{*r*}, *c*)**H**_{*i*}**P**(**1**_{*r*}, *c*)**H**_{*i*}}/ $\sqrt{rk(H_i)rk(H_i)}$ it holds that $\eta_r(\mathbf{K}) \ge b$, for b > 0 constant.

First, (A) imposes restrictions on θ_0 . It encompasses the conditions of positive covariance parameters and nondegenerativity as introduced by [14]. The condition $\max(\theta) / \min(\theta) \le c$ allows for all components of θ_0 to be either arbitrarily small or large, but not both. Condition (B) ensures that the covariance matrix is not singular and that θ_0 can be estimated consistently. Next, (C) corresponds to [8, Eq. (3.47)] and ensures that β_0 can be estimated consistently, which is the key advantage in the low-dimensional setting, that is for p < n. An extension to the case $n \le p$ is discussed in Section 7. Furthermore, (D) ensures that the normalizing sequence in Theorem 1 bounds the REML estimator for θ_0 uniformly. Finally, (E) relates to the estimator of the covariance parameters θ_0 . It encompasses the condition that $\mathbf{H}_k \in \mathbb{R}^{n \times n}$, $k \in \{1, ..., r\}$, are linearly independent, which corresponds to the condition of identifiability of covariance parameters from [14]. Since all involved matrices are known, the last two conditions are typically easy to verify, as the next two examples show.

Example 1. Consider a linear mixed model (1) with $\mathbf{X}_i = \mathbf{Z}_i = \mathbf{1}_{n_i}$, $\boldsymbol{\Psi}(\boldsymbol{\theta}_0) = \sigma_v^2$ and $\boldsymbol{\Omega}_i(\boldsymbol{\theta}_0) = \sigma_e^2 \mathbf{I}_{n_i}$, so that $\mathbf{V}(\boldsymbol{\theta}_0) = \sigma_v^2 \mathbf{H}_1 + \sigma_e^2 \mathbf{H}_2$, with $\mathbf{H}_1 = \text{blockdiag}(\mathbf{1}_{n_1}\mathbf{1}_{n_1}^\top, \dots, \mathbf{1}_{n_m}\mathbf{1}_{n_m}^\top)$, m > 1, $\mathbf{H}_2 = \mathbf{I}_n$ and $\boldsymbol{\theta}_0 = (\sigma_v^2, \sigma_e^2)^\top$. Here $m = \min_{k \in \{1, \dots, r\}} \operatorname{rk}(\mathbf{H}_k)$ translates to the number of clusters in the LMM. It is well known that the ratio of the entries in $\boldsymbol{\theta}_0$ is influential for its estimation, see [16]. In this LMM the intraclass correlation ratio

$$\gamma_i = \frac{\sigma_v^2}{\sigma_v^2 + \sigma_e^2/n_i}$$

measures the class variability relative to the total variability. If $\gamma_i \approx 0$, σ_v^2 cannot be estimated reliably. Condition (A) translates into a positive lower bound for γ_i . Further, $\mathbf{H}_1 \ge 0$ and $\mathbf{H}_2 > 0$, which corresponds to condition (B). Condition (C) is trivially satisfied. Next, (D) and (E) can be verified directly, since all quantities involved are known. For, e.g., c = 1, $n_1 = \cdots = n_m = 2$, this gives $\omega = 1/3$ and $\overline{\omega} = 2/3$ and

$$\mathbf{K} = \frac{1}{9} \begin{pmatrix} 4\frac{m-1}{m} & \sqrt{2}\frac{m-1}{m} \\ \sqrt{2}\frac{m-1}{m} & \frac{10m-1}{2m} \end{pmatrix},$$

for which b = 1/5. Note that condition (D) includes cases in which n_i is bounded as well as cases where n_i is unbounded, $i \in \{1, ..., m\}$. This has implications at which rate θ_0 and in turn β_0 are estimated. For instance, the rate in which σ_v^2 is estimated via REML is governed by m, rather than the total number of observations $n = \sum_{i=1}^m n_i$. Clearly, if all n_i , $i \in \{1, ..., m\}$ are bounded, m = O(n). Since (D) holds for both asymptotic scenarios and for balanced $(n_1 = \cdots = n_m)$ as well as unbalanced $(n_i \neq n_j)$ LMMs, the condition is sufficiently general for most applications.

Example 2. Eq. (2) allows for more general covariance structures than (1) in which the observations are not necessarily clustered as independent blocks. Assume a model with two random effects $\mathbf{v} \in \mathbb{R}^{\hat{m}}$ and $\tilde{\mathbf{v}} \in \mathbb{R}^{\hat{m}}$

$$\mathbf{y} = \mathbf{X}\boldsymbol{\beta}_0 + \mathbf{Z}\mathbf{v} + \mathbf{Z}\tilde{\mathbf{v}} + \boldsymbol{\varepsilon},$$

where $\mathbf{Z} = \text{blockdiag}(\mathbf{1}_{n_i})_{i=1}^m$, $\tilde{\mathbf{Z}} = \text{blockdiag}(\mathbf{1}_{\tilde{n}_j})_{j=1}^{\tilde{m}}$. Condition (B) is fulfilled if $\mathbf{H}_1 = \mathbf{Z}\mathbf{Z}^{\top}$ and $\mathbf{H}_2 = \tilde{\mathbf{Z}}\tilde{\mathbf{Z}}^{\top}$. However, if $m = \tilde{m}$ and $n_i = \tilde{n}_i$ for all $i \in \{1, ..., m\}$, (E) is not satisfied. A variation of such a model is discussed in Section 6.

3. Uniform Cramér consistency of the covariance parameters

The key idea in deriving uniformly valid confidence sets in LMMs is that the estimation of β_0 and θ_0 can be separated. The virtue of restricted maximum likelihood (REML) lies in estimating the covariance parameters, while accounting for the loss of degrees of freedom from estimation of β_0 [36]. The REML estimator for θ_0 is a maximizer of

$$\ell_{R}(\boldsymbol{\theta}) = -\frac{1}{2} \ln |\mathbf{V}(\boldsymbol{\theta})| - \frac{1}{2} \ln |\mathbf{X}^{\top} \mathbf{V}(\boldsymbol{\theta})^{-1} \mathbf{X}| - \frac{1}{2} \mathbf{y}^{\top} \mathbf{P}(\boldsymbol{\theta}) \mathbf{y},$$
(5)

where $\mathbf{P}(\boldsymbol{\theta}) = \mathbf{P}(\boldsymbol{\theta}, 1)$, as defined in (E) [17]. By construction, ℓ_R is constant with respect to $\boldsymbol{\beta}_0$ as $\mathbf{P}(\boldsymbol{\theta})\mathbf{X}\boldsymbol{\beta}_0 = \mathbf{0}_n$.

As interest lies in uniformly valid inference, uniform consistency of the REML estimator for θ_0 is required. Consistency of general maximum likelihood estimators (MLE) was first shown by [40], and uniform MLE consistency by [24]. Both required the parameter space to be compact, as well as independently drawn observations. A consistency (but not uniform consistency) result that relaxes those assumptions was first given by [41,42]. [23] extended Weiss's results for MLEs in LMMs, while [6,7] did so for REML estimators. [14] considered the REML estimator under non-normal LMMs and for unbounded *p*. However, to the best of our knowledge, uniform consistency of REML estimators in LMMs has not been treated.

In the following theorem, we prove the Cramér-type uniform consistency over Θ for REML estimators in LMMs. That is, the uniform consistency is shown for a sequence of local maximizers of (5), but this sequence does not have to be unique. In fact, for unbalanced panels in LMMs, that is if $n_i \neq n_j$, for $i \neq j$ in (1), it cannot be guaranteed that a maximizer of (5) is unique [15].

Theorem 1. Let model (2) and (A)–(E) hold and let $\nu_k(\theta_0) = \operatorname{rk}(\mathbf{H}_k)^{-1/2}\operatorname{tr}\{\mathbf{V}(\theta_0)^{-1}\mathbf{H}_k\}$ for $k \in \{1, \ldots, r\}$ and $m = \min_{k \in \{1, \ldots, r\}} \{\operatorname{rk}(\mathbf{H}_k)\} \to \infty$. Then, there exists a sequence $\hat{\theta}$ of local maximizers of (5), such that

$$\nu_k(\boldsymbol{\theta}_0) \left| \hat{\theta}_k - \theta_{0,k} \right| = O_P$$

uniformly over $\theta_0 \in \Theta$.

Throughout the paper, the notation $Z_m = O_P(a)$ for a sequence of random variables Z_m , $m \in \mathbb{N}$, is understood to hold uniformly. More concretely, it is defined that for all $\varepsilon > 0$, there exist constants M_{ε} , N_{ε} such that for all $m > N_{\varepsilon}$: $\inf_{\theta_0 \in \Theta} \Pr_{\theta_0}(Z_m \leq M_{\varepsilon}) \geq 1 - \varepsilon$. Regarding the normalizing sequence $v_k(\theta_0)$, note that $-E_{\theta_0}\{\partial^2 \ell_R(\theta)/\partial \theta_k^2|_{\theta_0}\} = O\{v_k(\theta_0)^2\}$, see (14) for details. The proof of Theorem 1 follows the lines of [24,41]. We extend their reasoning to account for a uniformity argument over Θ by the construction of the normalizing sequence $v_k(\theta_0)$. If θ_0 were fixed, the normalizing sequence would reduce to $\sqrt{rk(\mathbf{H}_k)}$, which corresponds to the parametric rate in classical linear models [14]. However, the specific choice of $v_k(\theta_0)$ allows to prove uniform consistency with the help of the auxiliary result given in Lemma 1 in Section 8.

4. Uniform confidence regions for the coefficient parameters

Now we can derive a uniform confidence region for the parameter β_0 , based on the Lasso estimator. For linear models, it is known that the minimal (infimal) coverage for Lasso-based confidence sets over the space of coefficient parameters occurs when the parameter values are large in absolute value, depending only on the sign of the entries of β_0 . This finding carries over to infimal coverage over the space of coefficient and covariance parameters, as the latter, when estimated with REML, do not depend on the fixed effects.

Proposition 1. Let model (2) and (A)–(E) hold. Then there exists a sequence $\hat{\theta}$ of local maximizers of (5) such that

$$\inf_{\substack{\boldsymbol{\beta}_{0}\in\mathbb{R}^{p}\\\boldsymbol{\theta}_{0}\in\Theta}}\Pr_{\boldsymbol{\beta}_{0},\boldsymbol{\theta}_{0}}\left\{\sqrt{n}\left(\hat{\boldsymbol{\beta}}_{L}-\boldsymbol{\beta}_{0}\right)\in E\left(\hat{\mathbf{C}},t\right)\right\}=\inf_{\substack{\mathbf{d}\in\{-1,1\}^{p}\\\boldsymbol{\theta}_{0}\in\Theta}}\Pr_{\boldsymbol{\theta}_{0}}\left\{\hat{\mathbf{u}}_{\mathbf{d}}\in E\left(\hat{\mathbf{C}},t\right)\right\}$$

The proof follows from [9, Theorem 1] as $\hat{\theta}$ is independent from β_0 by construction. Instead of evaluating the infimum for β_0 directly, the result states that a much simpler minimization over a discrete set is sufficient to evaluate the probability. In the context of LMMs, the key consequence of Proposition 1 is that, due to the orthogonality of $\hat{\theta}$ and β_0 , the uniform coverage probability can be separately evaluated for β_0 and θ_0 . For the latter, Theorem 1 allows to replace $\hat{\theta}$ by θ_0 at a cost of an error term with a tractable dependency on θ_0 . Eventually, this leads to the following result.

Theorem 2. Let model (2) and (A)–(E) hold and $m = \min_{k \in \{1,...,r\}} \{ \operatorname{rk}(\mathbf{H}_k) \} \to \infty$. Then, there exists a sequence $\hat{\theta}$ of local maximizers of (5) such that

$$\hat{\tau} = \max_{\mathbf{d} \in \{-1,1\}^p} \chi^2_{p,1-\alpha} \left(\left\| \hat{\mathbf{C}}^{-1/2} \boldsymbol{\Lambda} \mathbf{d} \right\|^2 \right)$$

the quantile of the non-central χ_p^2 -distribution, and it holds for all $\theta_0 \in \Theta$ that

$$\inf_{\substack{\boldsymbol{\beta}_0 \in \mathbb{R}^p\\\boldsymbol{\theta}_0 \in \Theta}} \Pr_{\boldsymbol{\beta}_0, \boldsymbol{\theta}_0} \left\{ \sqrt{n} \left(\hat{\boldsymbol{\beta}}_L - \boldsymbol{\beta}_0 \right) \in E\left(\hat{\mathbf{C}}, \hat{\tau} \right) \right\} = 1 - \alpha + O\left(\frac{1}{\sqrt{m}} \right).$$

The result differs from Proposition 1 in two aspects. First, the minimization over $\mathbf{d} \in \{-1, 1\}^p$ has been shifted to the choice of the non-centrality parameter of the χ^2 -distribution, based on Propositions 4 and 5 from [9]. Second, the estimation of θ_0 only contributes to an error term of uniformly vanishing, parametric order $m^{-1/2}$. The rate is given by the *m* observations that contribute to the estimation of $\theta_{0,k'}$, $k' = \operatorname{argmin}_{k \in \{1,...,r\}} \operatorname{rk}(\mathbf{H}_k)$. The crucial finding is that the error terms that occur in the estimation of both $\boldsymbol{\beta}_0$ and θ_0 and which depend on $\boldsymbol{\theta}_0$ eventually translate to an error term uniformly independent of θ_0 . As a consequence of this result, it follows that the Lasso-based confidence ellipsoid

$$M = \left\{ \boldsymbol{\beta} \in \mathbb{R}^{p} : n \left\| \hat{\mathbf{C}}^{1/2} \left(\hat{\boldsymbol{\beta}}_{L} - \boldsymbol{\beta} \right) \right\|^{2} \leq \hat{\tau} \right\}$$
(6)

attains nominal coverage uniformly up to an error of a parametric rate. By separating the estimation of β_0 with penalization and θ_0 without it, the result allows to infer about the fixed effects for data with a complex dependency structure. From the orthogonality argument, it is clear that the penalization on β_0 does not interfere with inference based on $\hat{\theta}$, for which existing theory can be applied. Since the REML estimates exhibit no analytically tractable finite sample distribution, the approximation with a χ^2 -quantile is only valid asymptotically. Although this implies that the resulting testing procedure is not of nominal level $1 - \alpha$ as discussed in [19], the simulation in Section 5 suggests that this error term has little influence in finite samples. Most importantly, however, is that the error term does not depend on β_0 and θ_0 and the stated coverage probability is attained uniformly over both the space of regression coefficients and covariance parameters. Nevertheless, the precise coverage probability depends on the sign of the entries of β_0 . Specifically, all but two orthants exhibit an overcoverage. This effect is visualized and discussed in Fig. 1 in Section 5.

When compared to standard confidence sets based on estimation procedures without variable selection, it is wellknown that the confidence set *M* is larger [31]. In many applications, however, smaller models and thus selection of covariates is preferred. In such cases, $\hat{\beta}_L$ and inference based upon it offer an alternative, despite the Lasso's difficult distributional properties.

5. Simulations

In this section, the finite sample coverage probabilities of the confidence set (6) are compared with several alternatives. First, with a least-squares-based approach, which offers uniform coverage but no selection. Second, with a least-squaresbased approach after selecting a model with AIC, resulting in type-I-error inflation. And finally, we demonstrate that a least-squares-based confidence set centered at $\hat{\beta}_L$ does not meet the nominal level. For visualization, we use the following simulation design for two coefficient parameters. Consider the 'random intercept model', a special case of the model (1),

$$\mathbf{y}_{ij} = \mathbf{x}_{ij}^{\top} \boldsymbol{\beta}_0 + v_i + u_{ij}, \quad i \in \{1, \dots, m\}, \quad j \in \{1, \dots, n_i\}, \quad u_{ij} \stackrel{i.i.d.}{\sim} \mathcal{N}(0, \sigma_u^2), \quad v_i \stackrel{i.i.d.}{\sim} \mathcal{N}(0, \sigma_v^2).$$
(7)

We fix the parameters to m = 20, n = 400, $n_i = 20$, $\sigma_u = \sigma_v = 4$. Note that the parameter m that determines the convergence rate is quite small. For visualization purposes, we restrict the simulation to p = 2 parameters. The tuning parameters are chosen to be $\lambda_i = n^{1/2}/2$ for $i \in \{1, 2\}$, which corresponds to a conservative tuning regime. The entries $\mathbf{x}_{ij} = (x_{1ij}, x_{2ij})^{\top}$ of the matrix of covariables are independently drawn once from $\mathcal{N}(0, 4)$ so that the fixed and random effects are of a comparable magnitude. For each $\boldsymbol{\beta}_0 \in [-4, 4] \times [-4, 4]$ on an equidistant grid of 81×81 , 2, 000 Monte Carlo simulations were carried out. For each simulation, the empirical coverage probability is computed by checking if $\boldsymbol{\beta}_0 \in M$, for M with $\alpha = 0.05$ from (6). Fig. 1 shows the results.



Fig. 1. Only confidence sets based on the Lasso as in (6) or WLS estimator (left column) achieve a nominal level over the whole parameter space. The former is conservative around the origin and axes. Naïvely applying WLS sets to the Lasso or WLS estimator after AIC variable selection (right column) yields undercoverage.

The probabilities shown are average empirical coverages for all configurations of β_0 . First consider the classical confidence sets based on the weighted least-squares (WLS) estimator $\hat{\beta}_{WLS} = \{\mathbf{X}^\top \mathbf{V}(\hat{\theta})^{-1}\mathbf{X}\}^{-1}\mathbf{X}^\top \mathbf{V}(\hat{\theta})^{-1}\mathbf{y}$ (bottom left), with no variable selection involved. As the distribution of $\hat{\beta}_{WLS} - \beta_0$ is independent of β_0 , the coverage based on its confidence set is attained uniformly over the space of coefficient parameters. One finds that no deviations from the nominal level of 95% can be observed in the simulation. Next, consider the confidence set based on the Lasso as given in (6) (upper left). No undercoverage is observed over the entire parameter space. Nominal coverage, up to an error of order $O(m^{-1/2})$ due to the estimation of random effects, is attained for $\operatorname{sign}(\beta_1) = -\operatorname{sign}(\beta_2)$. The other orthants exhibit a slight overcoverage (light blue), whereas a significant overcoverage (white) occurs at the axes and around the origin. The latter effects are due to the event of variable selection when a component in $\hat{\beta}_L$ is set to zero. Hence, at the axes, a coverage close to 100% is achieved, and the 95%-confidence sets prove to be too wide. These findings are in line with the example of the linear regression model from [9, Fig. 4].

The overcoverage in all but two orthants is a consequence of Lemma 4. The confidence sets are based on the minimizers in (4), which depend on **d**, the signs of the coefficient parameters. The nominal coverage probability is attained at the orthants specified by $\mathbf{d}^* = \operatorname{argmax}_{\mathbf{d} \in \{-1,1\}^p} \|\mathbf{C}^{-1/2} \Lambda \mathbf{d}\|^2$, corresponding to the upper left and lower right orthants in Fig. 1 (upper left). In these, the coverage probabilities vary between 93.4% and 95.8%, which is close to the nominal level of 95%. This indicates that the additional uncertainty induced by the estimation of θ_0 is not substantial even if *m* is small. If the coefficient signs of β_1 and β_2 differ from \mathbf{d}^* , the corresponding confidence sets are conservative. In Fig. 1 (upper left), the coverage probabilities vary between 94.6% and 96.8% in such orthants, which is close to the nominal level, too.

In contrast to the methods on the left column in Fig. 1, which meet the nominal level thoroughly, but without variable selection (bottom left) or conservatively, but with selection (upper left), two additional naïve approaches are displayed on the right column. First, one observes that naïvely applying classical WLS confidence sets around a WLS estimator

 $\hat{\beta}_{WLS}$ after performing variable selection with AIC (bottom right) yields inconsistent coverages over the parameter space. Type-I-error inflation [1] occurs in regions with undercoverage (red) in the event of variable selection at the axes. Another approach is applying a WLS confidence set around the estimator $\hat{\beta}_L$ (upper right). Again, these sets are not theoretically justified and indeed, an overcoverage occurs at the origin, whereas a severe undercoverage over the rest of the coefficient parameter space. Both naïve approaches thus yield misleading confidence sets, and their use is inadvisable.

We have also experimented with more complex covariance matrices, e.g., taking serially dependent observations within a cluster. While this increases the computational burden, the overall conclusion remains the same. Similarly, if more than two covariates are considered (p > 2), the confidence sets will meet nominal level in two out of 2^p orthants, and show a slight overcoverage in the remaining 2^{p-1} . If β_0 has close to zero entries, the confidence sets are very conservative, and the coverage probabilities close to one.

6. Application

We demonstrate the performance of the confidence sets based on the Lasso and WLS with no variable selection on a data set on the ecological status of lakes in the northeastern United States collected by the [38]. The set has been thoroughly investigated in previous studies [3,26]. The model of interest is given by

$$\mathbf{y} = \mathbf{1}_n \beta_1 + \mathbf{x} \beta_2 + \mathbf{Z} \mathbf{v} + \mathbf{D} \mathbf{u} + \boldsymbol{\epsilon}, \quad \mathbf{v} \sim \mathcal{N}(\mathbf{0}_m, \sigma_v^2 \mathbf{I}_m), \quad \mathbf{u} \sim \mathcal{N}(\mathbf{0}_K, \sigma_u^2 \mathbf{I}_K), \quad \boldsymbol{\epsilon} \sim \mathcal{N}(\mathbf{0}_n, \sigma_e^2 \mathbf{I}_n).$$

The response variable of interest is the acid-neutralizing capacity in equivalents per liter, which indicates the lake's environmental state. It is related to a design matrix that includes an intercept and the respective elevation in meters. In total, the n = 551 lakes are partitioned via m = 19 hydrological unit codes with $1 \le n_i \le 13$, which enter the model as random effect intercept, i.e., $\mathbf{Z} = \text{blockdiag}(\mathbf{1}_{n_i})_{i=1}^n$. Finally, a penalized spline with K = 80 knots accounts for spatial proximity effects. It is included as a second random effect, for which \mathbf{D} is a transformed radial basis as described by [26].

The penalization parameters for the Lasso are set to $\lambda_1 = \lambda_2 = 16.9$ by leave-one-out- cross-validation obtained with the R-function glmnet. This value is close to $\sqrt{n}/2 \approx 11.74$, which corresponds to a conservative tuning regime. The fitted model does not indicate any deviation from regularity conditions (A)–(E). In particular, $\max(\hat{\theta}) / \min(\hat{\theta}) \approx 9.99$. The estimated fixed effects are given as

$$\hat{\boldsymbol{\beta}}_{L} = (0, -0.0010)^{\top}, \quad \hat{\boldsymbol{\beta}}_{WLS} = (-0.5589, -0.0011)^{\top}$$

In previous studies, the intercept has been deemed insignificant [26]. Indeed, the estimated standard deviations are given by $s(\hat{\beta}_{WLS,1}) = 2.2130$ and $s(\hat{\beta}_{WLS,2}) = 0.0001$. Consequently, the Lasso estimator sets the intercept equal to zero. This corresponds to a selection event on the axes in Fig. 1 (upper left). Such suffers from severe overcoverage. A 99%-confidence set M_L based on the Lasso is given by M from (6), with $\hat{\tau} = \chi^2_{2,0.99}(1398) \approx 1578$. The WLS based set M_{WLS} is given as M, but $\hat{\beta}_L$ replaced with $\hat{\beta}_{WLS}$ and $\hat{\tau}$ with $\chi^2_{2,0.99} \approx 9$. The Lasso-based confidence ellipse is much larger than the WLSbased confidence ellipse. This does not indicate that the former are useless, however. Much of their additional volume is distributed along the axis of the non-included intercept.

The additional uncertainty induced by the Lasso in case of no selection event can be assessed by performing inference on β_2 only. This gives confidence intervals for the elevation coefficient

$$M_L(\beta_2) = [-14.80, -7.13] \times 10^{-4}, \quad M_{WLS}(\beta_2) = [-13.73, -6.06] \times 10^{-4}.$$

Both sets are constructed similarly. While for M_L , the corresponding quantile is from the non-central χ_1^2 -distribution with a non-centrality parameter close to zero, M_{WLS} is based upon a central χ_1^2 -quantile. Therefore, the respective interval lengths are almost equal, with M_L being narrowly wider by the sixth decimal place. However, both intervals differ in location. The Lasso based $M_L(\beta_2)$ is centered around $\hat{\beta}_{L,2}$ and thus covers a different part of the real line than the WLS based $M_{WLS}(\beta_2)$. Consequently, although both intervals are of almost equal length, inference based on the Lasso indeed differs from WLS-based inference, even in the case of no selection event. The conclusions of this analysis remain valid if tuning parameters are set to the theoretical value $\lambda_1 = \lambda_2 = \sqrt{n}/2 \approx 11.74$.

Altogether, this case example confirms that if interest lies in narrow confidence sets, WLS sets are smaller and should be employed. Simultaneously performing variable selection with the Lasso always evokes uncertainty. Nevertheless, confidence sets based on the Lasso are not always much larger than their WLS counterparts. When smaller models are preferable, those sets can be useful for assessing uncertainty and hypothesis testing.

7. Discussion

This contribution presents a solution for inference in a low-dimensional LMM in which the fixed effects are estimated with a Lasso-type penalization. The uniformly valid confidence sets for the fixed effects based on a Lasso estimator, which performs estimation and model selection simultaneously, have been constructed. We suggest a two-stage estimation procedure, where the covariance parameters are estimated via a REML estimator $\hat{\theta}$ first, and the coefficient parameters β_0 with $\hat{\theta}$ plugged in second. In particular, this allows to treat regression coefficients and covariance parameters separately. Using this approach, we decisively simplify the verification of the theoretical properties of parameter estimators. Finally,

it can be shown that the resulting confidence sets are uniformly valid over both the coefficient and covariance parameters spaces.

It is well known that employing standard WLS-based confidence sets without model selection will always yield smaller confidence sets than post-selection inference [31]. However, selecting parsimonious models is often a key concern for practitioners. Although resulting post-selection confidence sets will include additional uncertainty due to variable selection, Sections 5 and 6 indicate that derived confidence sets are useful. Moreover, our results provide information about the joint estimation and variable selection uncertainty of the Lasso in the context of LMMs by quantifying the probability with which the estimation error $\hat{\beta}_L - \beta_0$ lies in the proposed ellipse.

The novelty of our method is that, to the best of our knowledge, this work is the first that considers uniform inference based on the Lasso for an elaborate covariance structure, as given in LMMs. Another new important theoretical result of this article is the proof of uniform consistency of Cramér type for the REML estimators of the covariance parameters. We expect that our approach can serve as a basis for proper inference for an estimation procedure that penalizes both fixed and random effects.

The derived confidence regions are based on the Lasso selection procedure, but are valid regardless of the selected model. In this, our methodology differs from selective inference methods for LMMs as proposed by [34].

It is important to note that the proposed method does not account for any stochastic selection of the tuning parameter λ . Instead, the result in Theorem 2 is a finite sample result that holds for any a-priori choice of λ . Setting $\lambda = n^{1/2}/2$ as in Section 5 ensures conservative model selection and uniformly valid inference. If λ is estimated or determined with any data-driven procedure, it emits additional volatility that has to be quantified. This holds a fortiori for procedures such as the adaptive Lasso where even the penalty term itself is random. While the penalty term of the smoothly clipped absolute deviations estimator (SCAD) [10] does not exhibit this property, it is particularly intricate to treat and leads to a non-convex objective function. Therefore, while these methods behave similarly to the ordinary Lasso, an extension of our findings for them requires substantial additional research, which is out of the scope of the current work. It should be noted that among Lasso, adaptive Lasso, and SCAD, [33] show that in the case of orthogonal regressors, using the Lasso leads to the shortest intervals in finite samples, strengthening our approach.

In high dimensions ($p \ge n$), confidence sets in connection with the Lasso penalization have been addressed, e.g., by the de-biasing approach of [39] and related works. For such settings, the ansatz of [9] is not applicable as the matrix **C** is not invertible. Moreover, the classical REML methodology is defined in a low-dimensional setting only, as θ_0 is estimated by the data projected onto the space orthogonal to the columns of **X**. Extensions for REML to higher dimensions lose the property that $\hat{\theta}$ is independent of β_0 [35].

8. Proofs

To prove Theorem 1, we first derive the following lemma.

Lemma 1. Let conditions (A)–(E) hold, $m = \min\{rk(\mathbf{H}_1), \ldots, rk(\mathbf{H}_r)\}$ and let K be a finite set drawn with replacement from $\{1, \ldots, r\}$ with |K| > 1. Then, for any $\theta \in \Theta$,

$$\operatorname{tr}\left\{\prod_{k\in K}\mathbf{P}(\boldsymbol{\theta})\mathbf{H}_{k}\right\} \leq \left(\frac{c}{\underline{\omega}}\right)^{|K|-1} m^{1-|K|/2} \prod_{k\in K} \nu_{k}(\boldsymbol{\theta}).$$
(8)

The result is shown using that the eigenvalues of the matrices of interest are smaller or equal to one and subsequently employing regularity condition (D).

Proof. Consider the symmetric and positive semi-definite matrices

$$\mathbf{A}_{k} = \mathbf{A}_{k}(\boldsymbol{\theta}) = \mathbf{V}(\boldsymbol{\theta})^{-1/2} \mathbf{H}_{k} \mathbf{V}(\boldsymbol{\theta})^{-1/2}, \quad k \in \{1, \dots, r\},$$

$$\mathbf{B} = \mathbf{B}(\boldsymbol{\theta}) = \mathbf{I}_{n} - \mathbf{V}(\boldsymbol{\theta})^{-1/2} \mathbf{X} \{\mathbf{X}^{\top} \mathbf{V}(\boldsymbol{\theta})^{-1} \mathbf{X}\}^{-1} \mathbf{X}^{\top} \mathbf{V}(\boldsymbol{\theta})^{-1/2},$$

(9)

so that $tr\{\prod_{k\in K} \mathbf{P}(\boldsymbol{\theta})\mathbf{H}_k\} = tr\{\prod_{k\in K} \mathbf{A}_k\mathbf{B}\}$. Since **B** is a projection matrix with eigenvalues zero or one, proceeding as in the proof of Theorem 1.1 (Hölder inequality) from [22] gives

$$\operatorname{tr}\left(\prod_{k\in K}\mathbf{A}_{k}\mathbf{B}\right) \leq \sum_{i=1}^{n}\prod_{k\in K}\left\{\eta_{i}(\mathbf{A}_{k})\eta_{i}(\mathbf{B})\right\} \leq \sum_{i=1}^{n}\prod_{k\in K}\eta_{i}(\mathbf{A}_{k}) \leq \prod_{k\in K}\operatorname{tr}\left(\mathbf{A}_{k}^{|K|}\right)^{1/|K|}.$$
(10)

We continue by evaluating tr($\mathbf{A}_{k}^{|K|}$). First note that conditions (A) and (D) imply

$$c^{-1}\underline{\omega} \le \theta_k \frac{\operatorname{tr}(\mathbf{A}_k)}{\operatorname{rk}(\mathbf{H}_k)} \le c \ \overline{\omega}.$$
(11)

For k = r, let $\eta'_i = \eta_i (\mathbf{H}_r^{-1} \sum_{l=1}^{r-1} \theta_l / \theta_r \mathbf{H}_l) \ge 0$. Then, the above inequality yields

$$\operatorname{tr}\left(\mathbf{A}_{r}^{|K|}\right) = \frac{1}{\theta_{r}^{|K|}} \operatorname{tr}\left\{ \left(\mathbf{I}_{n} + \mathbf{H}_{r}^{-1} \sum_{l=1}^{r-1} \frac{\theta_{l}}{\theta_{r}} \mathbf{H}_{k}\right)^{-|K|} \right\} = \frac{1}{\theta_{r}^{|K|}} \sum_{i=1}^{n} \frac{1}{(1+\eta_{i}')^{|K|}} \leq \frac{1}{\theta_{r}^{|K|}} \sum_{i=1}^{n} \frac{1}{1+\eta_{i}'} = \frac{\operatorname{tr}(\mathbf{A}_{r})}{\theta_{r}^{|K|-1}} \\ \leq \left(\frac{c}{\omega}\right)^{|K|-1} \frac{\operatorname{tr}(\mathbf{A}_{r})^{|K|}}{\operatorname{rk}(\mathbf{H}_{r})^{|K|-1}},$$

and, similarly for $k \in \{1, ..., r-1\}$, let $\eta_i'' = \eta_i \{ \left(\sum_{l \neq k} \theta_l / \theta_k \mathbf{H}_l \right)^{-1} \mathbf{H}_k \} \ge 0$. This gives

$$\operatorname{tr}(\mathbf{A}_{k}^{|K|}) = \frac{1}{\theta_{k}^{|K|}} \operatorname{tr}\left\{ \left(\mathbf{I}_{n} - \left[\mathbf{I}_{n} + \left(\sum_{l \neq k} \frac{\theta_{l}}{\theta_{k}} \mathbf{H}_{l} \right)^{-1} \mathbf{H}_{k} \right]^{-1} \right)^{|K|} \right\} = \frac{1}{\theta_{k}^{|K|}} \sum_{i=1}^{n} \left(\frac{\eta_{i}^{''}}{1 + \eta_{i}^{''}} \right)^{|K|}$$

$$\leq \frac{1}{\theta_{k}^{|K|}} \sum_{i=1}^{n} \frac{\eta_{i}^{''}}{1 + \eta_{i}^{''}} = \frac{\operatorname{tr}(\mathbf{A}_{k})}{\theta_{k}^{|K|-1}} \leq \left(\frac{c}{\underline{\omega}} \right)^{|K|-1} \frac{\operatorname{tr}(\mathbf{A}_{k})^{|K|}}{\operatorname{tr}(\mathbf{H}_{k})^{|K|-1}}.$$

Altogether, for $k \in \{1, \ldots, r\}$, this gives

$$\operatorname{tr}(\mathbf{A}_{k}^{|K|}) \leq \frac{c^{|K|-1}}{\underline{\omega}^{|K|-1}} \frac{\operatorname{tr}(\mathbf{A}_{k})^{|K|}}{\operatorname{rk}(\mathbf{H}_{k})^{|K|-1}}.$$
(12)

Plugging this into (10) yields

$$\prod_{k\in K} \operatorname{tr} \left(\mathbf{A}_{k}^{|K|} \right)^{1/|K|} \leq \prod_{k\in K} \left(\frac{c}{\underline{\omega}} \right)^{\frac{|K|-1}{|K|}} \frac{\nu_{k}(\boldsymbol{\theta})}{\operatorname{rk}(\mathbf{H}_{k})^{\frac{|K|-2}{2|K|}}} \leq \left(\frac{c}{\underline{\omega}} \right)^{|K|-1} m^{1-|K|/2} \prod_{k\in K} \nu_{k}(\boldsymbol{\theta}),$$

which gives the claim. \Box

Note that under the conditions of Lemma 1 but with |K| = 1, (10) implies that

 $\operatorname{tr} \left\{ \mathbf{P}(\boldsymbol{\theta}) \mathbf{H}_k \right\} \le \operatorname{tr} \left\{ \mathbf{A}_k(\boldsymbol{\theta}) \right\}.$ (13)

Before getting into the proof of Theorem 1, we denote from now on $\mathbf{Q}_i(\theta) = \mathbf{P}(\theta)\mathbf{H}_i$, $\mathbf{Q}_{ij}(\theta) = \prod_{l \in \{i,j\}} \mathbf{P}(\theta)\mathbf{H}_l$, $\mathbf{Q}_{ijk}(\theta) = \prod_{l \in \{i,j\}} \mathbf{P}(\theta)\mathbf{H}_l$ to facilitate notation. In particular, Lemma 1 implies that

$$- E_{\boldsymbol{\theta}_0} \left\{ \frac{\partial^2 \ell_R}{\partial \theta_i^2}(\boldsymbol{\theta}) \bigg|_{\boldsymbol{\theta}_0} \right\} = \frac{1}{2} \operatorname{tr} \left\{ \mathbf{Q}_{ii}(\boldsymbol{\theta}_0) \right\} = O\{\nu_i(\boldsymbol{\theta}_0)^2\}.$$
(14)

It is straightforward to adapt Lemma 1 to the case in which $\mathbf{P}(\boldsymbol{\theta})$ is replaced by $\mathbf{V}(\boldsymbol{\theta})^{-1}$:

Corollary 1. Let model (2) and (A)–(E) hold and let K be a set drawn with replacement from $\{1, ..., r\}$ with |K| > 1. Then, for $\theta \in \Theta$,

$$\operatorname{tr}\left\{\prod_{k\in K}\mathbf{V}(\boldsymbol{\theta})^{-1}\mathbf{H}_{i}\right\} \leq \left(\frac{c}{\underline{\omega}}\right)^{|K|-1}\prod_{k\in K}\frac{\nu_{i}(\boldsymbol{\theta})}{\operatorname{rk}(\mathbf{H}_{i})^{\frac{|K|-2}{2|K|}}}.$$
(15)

Above results are related to the reverse inequality from [12, p. 232, Problem 4.1.P13], namely that for a positive definite matrix $\mathbf{Z} \in \mathbb{R}^{n \times n}$, $n \in \mathbb{N}$, it holds

$$\operatorname{tr}(\mathbf{Z}^2) \ge \frac{\operatorname{tr}(\mathbf{Z})^2}{\operatorname{rk}(\mathbf{Z})}.$$
(16)

Proof of Theorem 1. This proof is adapted to account for uniformity w.r.t. θ_0 from Lemma 7.2 (i) from [14] and closely follows the lines of [41]. The four boundedness properties below on the derivatives of the log-likelihood are surrogates of the boundedness conditions imposed on the log-likelihood directly by [24,40]. By a Taylor expansion, the conditions below also include the boundedness of the log-likelihood. They are further required to omit the compactness condition of Θ , by constructing a compact set Θ'_{ϵ} , see details below. For all $i, j \in \{1, ..., r\}$, it holds for any θ_0 that

(i)
$$E_{\theta_0} \left\{ \frac{\partial \ell_R}{\partial \theta_i}(\boldsymbol{\theta}) \Big|_{\theta_0} \right\} = 0,$$

(ii) $\frac{1}{\nu_i(\boldsymbol{\theta}_0)} \frac{\partial \ell_R}{\partial \theta_i}(\boldsymbol{\theta}) \Big|_{\theta_0} = O_P(1),$

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(iii) For
$$\mathbf{J}(\boldsymbol{\theta}_{0})$$
 with $J_{ij}(\boldsymbol{\theta}_{0}) = -\frac{1}{\nu_{i}(\boldsymbol{\theta}_{0})\nu_{j}(\boldsymbol{\theta}_{0})} \mathbf{E}_{\boldsymbol{\theta}_{0}} \left\{ \frac{\partial^{2}\ell_{R}}{\partial\theta_{i}\partial\theta_{j}}(\boldsymbol{\theta}) \Big|_{\boldsymbol{\theta}_{0}} \right\}$ it holds that $\eta_{r}\{\mathbf{J}(\boldsymbol{\theta}_{0})\} \ge c' > 0$, for some constant $c' > 0$.
(iv) $\zeta(\boldsymbol{\theta}, \boldsymbol{\theta}_{0}) = \frac{1}{\nu_{i}(\boldsymbol{\theta}_{0})\nu_{j}(\boldsymbol{\theta}_{0})} \frac{\partial^{2}\ell_{R}}{\partial\theta_{i}\partial\theta_{j}}(\boldsymbol{\theta}) + J_{ij}(\boldsymbol{\theta}_{0}) = O_{P}\left(\frac{q}{\sqrt{m}}\right)$ for $\boldsymbol{\theta} \in \Theta_{q} = \{\boldsymbol{\theta} : \nu_{i}(\boldsymbol{\theta}_{0})|\theta_{i} - \theta_{0,i}| \le q\}$,

where $q = m^{1/(6+\delta)}$ for some $\delta > 0$ and $m = \min_{k \in \{1,...,r\}} \operatorname{rk}(\mathbf{H}_i)$. For readability, suppress the dependency from θ when the argument is clear from the context. Now, (i)–(iv) are shown.

(i) Note that $\partial \mathbf{P} / \partial \theta_i = -\mathbf{P} \mathbf{H}_i \mathbf{P}$ and $\mathbf{P} \mathbf{V} \mathbf{P} = \mathbf{P}$, as well as

$$\frac{\partial \ell_R}{\partial \theta_i}\Big|_{\boldsymbol{\theta}_0} = \frac{1}{2} \mathbf{y}^\top \mathbf{P} \mathbf{H}_i \mathbf{P} \mathbf{y} - \frac{1}{2} \mathrm{tr} \left(\mathbf{P} \mathbf{H}_i \right).$$

The claim now follows after taking expectations.

(ii) As $E(\partial \ell_R / \partial \theta_i |_{\theta_0}) = 0$ by (i), Chebyshev's inequality can be applied, it holds uniformly and gives that for any $\epsilon > 0$ there exists k > 0, such that

$$\sup_{\boldsymbol{\theta}_{0}\in\Theta} \Pr_{\boldsymbol{\theta}_{0}} \left\{ \operatorname{Var}_{\boldsymbol{\theta}_{0}} \left(\frac{\partial \ell_{R}}{\partial \theta_{i}} \Big|_{\boldsymbol{\theta}_{0}} \right)^{-1/2} \left| \frac{\partial \ell_{R}}{\partial \theta_{i}} \Big|_{\boldsymbol{\theta}_{0}} \right| \geq k \right\} \leq \frac{1}{k^{2}} < \epsilon,$$

and thus

$$\frac{\partial \ell_R}{\partial \theta_i} \bigg|_{\theta_0} = O_P \left\{ \operatorname{Var}_{\theta_0} \left(\frac{\partial \ell_R}{\partial \theta_i} \bigg|_{\theta_0} \right)^{1/2} \right\} = O_P(\nu_i),$$

where the last equation follows by $\operatorname{Var}_{\theta_0}(\partial \ell_R / \partial \theta_i|_{\theta_0}) = -E_{\theta_0}\{\partial^2 \ell_R / \partial \theta_i^2|_{\theta_0}\}$ and (14).

(iii) By (11), for all $i, j \in \{1, ..., r\}$,

$$J_{ij} = \frac{\operatorname{tr}(\mathbf{Q}_{ij})}{2\nu_i\nu_j} \geq \frac{\theta_{0,i}\theta_{0,j}\operatorname{tr}(\mathbf{Q}_{ij})}{2c^2\overline{\omega}^2\sqrt{\operatorname{rk}(\mathbf{H}_i)\operatorname{rk}(\mathbf{H}_j)}}.$$

By a similar argument,

$$\begin{aligned} \theta_{0,i}\theta_{0,j}\mathrm{tr}(\mathbf{Q}_{ij}) &= \theta_{0,i}\theta_{0,j}\mathrm{tr}\big[\mathbf{V}(\theta_0)^{-1}[\mathbf{I}_n - \mathbf{X}\{\mathbf{X}^{\top}\mathbf{V}(\theta_0)^{-1}\mathbf{X}\}^{-1}\mathbf{X}^{\top}\mathbf{V}(\theta_0)^{-1}]\mathbf{H}_i\mathbf{P}(\theta_0)\mathbf{H}_j\big] \\ &\geq \theta_{0,j}\frac{\min(\theta)}{\max(\theta)}\mathrm{tr}\big[\mathbf{V}(\mathbf{1}_r)^{-1}[\mathbf{I}_n - \mathbf{X}\{\mathbf{X}^{\top}\mathbf{V}(\theta_0)^{-1}\mathbf{X}\}^{-1}\mathbf{X}^{\top}\mathbf{V}(\theta_0)^{-1}]\mathbf{H}_i\mathbf{P}(\theta_0)\mathbf{H}_j\big] \\ &\geq \frac{\theta_{0,j}}{c}\mathrm{tr}\big[\mathbf{V}(\mathbf{1}_r)^{-1}[\mathbf{I}_n - c\mathbf{X}\{\mathbf{X}^{\top}\mathbf{V}(\mathbf{1}_r)^{-1}\mathbf{X}\}^{-1}\mathbf{X}^{\top}\mathbf{V}(\mathbf{1}_r)^{-1}]\mathbf{H}_i\mathbf{P}(\theta_0)\mathbf{H}_j\big] \geq \frac{1}{c^2}\mathrm{tr}\{\mathbf{P}(\mathbf{1}_r, c)\mathbf{H}_i\mathbf{P}(\mathbf{1}_r, c)\mathbf{H}_j\} \end{aligned}$$

It follows that

$$J_{ij}(\boldsymbol{\theta}_0) \geq \frac{\mathrm{tr}\{\mathbf{P}(\mathbf{1}_r, c)\mathbf{H}_i\mathbf{P}(\mathbf{1}_r, c)\mathbf{H}_j\}}{2c^4\overline{\omega}^2\sqrt{\mathrm{rk}(\mathbf{H}_i)\mathrm{rk}(\mathbf{H}_j)}}.$$

Now, for $\mathbf{a} \in \mathbb{R}^r$ with $\mathbf{a}^\top \mathbf{a} = 1$ eigenvector to $\eta_r \{ \mathbf{J}(\boldsymbol{\theta}_0) \}$,

$$\eta_r\{\mathbf{J}(\boldsymbol{\theta}_0)\} = \mathbf{a}^\top \mathbf{J}(\boldsymbol{\theta}_0) \mathbf{a} = \sum_{i=1}^r \sum_{j=1}^r a_i a_j J_{ij}(\boldsymbol{\theta}_0) \ge \frac{\eta_r\{\mathbf{K}\}}{2c^4 \overline{\omega}^2} \ge \frac{b}{2c^4 \overline{\omega}^2} > 0.$$

(iv) First, note that for $\theta \in \Theta_q$, (11) gives

$$\frac{\underline{\omega}\sqrt{\mathrm{rk}(\mathbf{H}_i)}}{c\theta_i} \leq \nu_i(\boldsymbol{\theta}) \leq \frac{c\overline{\omega}\sqrt{\mathrm{rk}(\mathbf{H}_i)}}{\theta_i}.$$

For any $t \in [0, 1]$ and $\tilde{\theta} = \theta_0 + t(\theta - \theta_0)$ this implies with (D) and Corollary 1 that

$$\operatorname{tr}\left\{\mathbf{V}(\tilde{\boldsymbol{\theta}})^{-1}\mathbf{H}_{j}\mathbf{V}(\tilde{\boldsymbol{\theta}})^{-1}\mathbf{H}_{i}\right\} \leq \frac{c}{\theta_{0,j}\theta_{0,i}}\operatorname{tr}\left\{\mathbf{V}(\mathbf{1}_{r})^{-1}\mathbf{H}_{j}\mathbf{V}(\mathbf{1}_{r})^{-1}\mathbf{H}_{i}\right\} \leq \frac{c^{2}}{\underline{\omega}}\frac{\operatorname{tr}\left\{\mathbf{V}(\mathbf{1}_{r})^{-1}\mathbf{H}_{j}\right\}\operatorname{tr}\left\{\mathbf{V}(\mathbf{1}_{r})^{-1}\mathbf{H}_{i}\right\}}{\theta_{0,j}\theta_{0,i}\sqrt{\operatorname{rk}(\mathbf{H}_{j})\operatorname{rk}(\mathbf{H}_{i})}} \leq \frac{c^{2}\overline{\omega}^{2}}{\underline{\omega}}\nu_{i}(\boldsymbol{\theta})\nu_{j}(\boldsymbol{\theta}).$$

A Taylor expansion around θ_0 yields

$$\nu_{i}(\boldsymbol{\theta}) = \nu_{i}(\boldsymbol{\theta}_{0}) + \frac{1}{\sqrt{\mathrm{rk}(\mathbf{H}_{i})}} \sum_{j=1}^{r} (\theta_{j} - \theta_{0,j}) \mathrm{tr} \left\{ \mathbf{V}(\tilde{\boldsymbol{\theta}})^{-1} \mathbf{H}_{j} \mathbf{V}(\tilde{\boldsymbol{\theta}})^{-1} \mathbf{H}_{i} \right\} \leq \nu_{i}(\boldsymbol{\theta}_{0}) \left\{ 1 + \frac{\overline{\omega}^{2} c^{3}}{\underline{\omega}^{2} \sqrt{m}} \sum_{j=1}^{r} \nu_{j}(\boldsymbol{\theta})(\theta_{j} - \theta_{0,j}) \right\}$$

$$\leq \nu_{i}(\boldsymbol{\theta}_{0}) \left\{ 1 + \frac{r \overline{\omega}^{2} c^{3} q}{\underline{\omega}^{2} \sqrt{m}} \right\}.$$
(17)

Below, abbreviate the constant $c' = r\overline{\omega}^2 c^3 / \underline{\omega}^2$ and note that $q/\sqrt{m} \to 0$. This implies that

$$\boldsymbol{\theta} \in \Theta_q \quad \Rightarrow \quad \nu_i(\boldsymbol{\theta})|\theta_i - \theta_{0,i}| \le q \left(1 + c' \frac{q}{\sqrt{m}}\right). \tag{18}$$

Now, by (18), it follows for $\theta \in \Theta_q$:

$$\mathbf{V}(\boldsymbol{\theta}_0) = \mathbf{V}(\boldsymbol{\theta}) + \sum_{k=1}^r (\theta_{0,k} - \theta_k) \mathbf{H}_k \le \mathbf{V}(\boldsymbol{\theta}) + \sum_{k=1}^r \mathbf{H}_k \frac{q}{\nu_k(\boldsymbol{\theta})} \left(1 + c' \frac{q}{\sqrt{m}} \right).$$

With Lemma 1, this gives

$$\mathbf{E}_{\boldsymbol{\theta}_{0}} \left\{ -\frac{1}{\nu_{i}(\boldsymbol{\theta}_{0})\nu_{j}(\boldsymbol{\theta}_{0})} \frac{\partial^{2}\ell_{R}}{\partial\theta_{i}\partial\theta_{j}}(\boldsymbol{\theta}) \right\} = -\frac{\mathrm{tr}\{\mathbf{Q}_{ij}(\boldsymbol{\theta})\}/2 - \mathrm{tr}\left\{\mathbf{Q}_{ij}(\boldsymbol{\theta})\mathbf{P}(\boldsymbol{\theta})\mathbf{V}(\boldsymbol{\theta}_{0})\right\}}{\nu_{i}(\boldsymbol{\theta}_{0})\nu_{j}(\boldsymbol{\theta}_{0})} \\ \leq \frac{\mathrm{tr}\{\mathbf{Q}_{ij}(\boldsymbol{\theta})\}}{2\nu_{i}(\boldsymbol{\theta}_{0})\nu_{j}(\boldsymbol{\theta}_{0})} + \sum_{k=1}^{r}q\frac{\mathrm{tr}\{\mathbf{Q}_{ijk}(\boldsymbol{\theta})\}}{\nu_{i}(\boldsymbol{\theta})\nu_{j}(\boldsymbol{\theta})\nu_{k}(\boldsymbol{\theta})} \left(1 + c'\frac{q}{\sqrt{m}}\right)^{3} \\ = \frac{\mathrm{tr}\{\mathbf{Q}_{ij}(\boldsymbol{\theta})\}}{2\nu_{i}(\boldsymbol{\theta}_{0})\nu_{j}(\boldsymbol{\theta}_{0})} + O\left(\frac{q}{\sqrt{m}}\right),$$

for $m = \min_{k \in \{1,...,r\}} \operatorname{rk}(\mathbf{H}_i)$ and all $\boldsymbol{\theta} \in \Theta_q$. Similarly, now dropping the argument for all quantities depending on $\boldsymbol{\theta} \in \Theta_q$ for the sake of readability, i.e. $v_i = v_i(\boldsymbol{\theta})$,

$$\begin{aligned} \operatorname{Var}_{\boldsymbol{\theta}_{0}}\left\{-\frac{1}{\nu_{i}(\boldsymbol{\theta}_{0})\nu_{j}(\boldsymbol{\theta}_{0})}\frac{\partial^{2}\ell_{R}}{\partial\theta_{i}\partial\theta_{j}}\right\} &= 2\frac{\operatorname{tr}\left\{\mathbf{Q}_{ij}\mathbf{PV}(\boldsymbol{\theta}_{0})\mathbf{Q}_{ij}\mathbf{PV}(\boldsymbol{\theta}_{0})\right\}}{\nu_{i}(\boldsymbol{\theta}_{0})^{2}\nu_{j}(\boldsymbol{\theta}_{0})^{2}} + O\left(\frac{q^{2}}{m}\right) \\ &= O\left\{\frac{\operatorname{tr}(\mathbf{Q}_{ij}\mathbf{Q}_{ij})}{\nu_{i}^{2}\nu_{j}^{2}} + 2q\sum_{k=1}^{r}\frac{\operatorname{tr}(\mathbf{Q}_{ij}\mathbf{Q}_{ijk})}{\nu_{i}^{2}\nu_{j}^{2}\nu_{k}} + q^{2}\sum_{k,l=1}^{r}\frac{\operatorname{tr}(\mathbf{Q}_{ijk}\mathbf{Q}_{ijl})}{\nu_{i}^{2}\nu_{j}^{2}\nu_{l}\nu_{k}} + \frac{q^{2}}{m}\right\} \\ &= O\left(\frac{1}{m^{1/2}} + \frac{2q}{m^{3/2}} + \frac{q^{2}}{m^{2}} + \frac{q^{2}}{m}\right) = O\left(\frac{q^{2}}{m}\right).\end{aligned}$$

Putting the previous two results together and proceeding as in (ii) with Chebyshev's inequality, this gives that for any θ_0 it holds

$$-\frac{1}{\nu_i(\boldsymbol{\theta}_0)\nu_j(\boldsymbol{\theta}_0)}\frac{\partial^2 \ell_R}{\partial \theta_i \partial \theta_j}(\boldsymbol{\theta}) = \frac{\mathrm{tr}\{\mathbf{Q}_{ij}(\boldsymbol{\theta})\}}{2\nu_i(\boldsymbol{\theta}_0)\nu_j(\boldsymbol{\theta}_0)} + O_P\left(\frac{q}{\sqrt{m}}\right)$$

for $\theta \in \Theta_q$. To prove (iv), this must hold for θ_0 on the right-hand side. Again, let $t \in [0, 1]$ and $\tilde{\theta} = \theta_0 + t(\theta - \theta_0)$. A Taylor expansion of tr{ $\mathbf{Q}_{ij}(\theta)$ } around tr{ $\mathbf{Q}_{ij}(\theta_0)$ } gives

$$\frac{\operatorname{tr}\{\mathbf{Q}_{ij}(\boldsymbol{\theta})\}}{2\nu_{i}(\boldsymbol{\theta}_{0})\nu_{j}(\boldsymbol{\theta}_{0})} = \frac{\operatorname{tr}\{\mathbf{Q}_{ij}(\boldsymbol{\theta}_{0})\}}{2\nu_{i}(\boldsymbol{\theta}_{0})\nu_{j}(\boldsymbol{\theta}_{0})} + \sum_{k=1}^{r} (\theta_{0,k} - \theta_{k}) \frac{\operatorname{tr}\{\mathbf{Q}_{ijk}(\tilde{\boldsymbol{\theta}}) + \mathbf{Q}_{ikj}(\tilde{\boldsymbol{\theta}})\}}{\nu_{i}(\boldsymbol{\theta}_{0})\nu_{j}(\boldsymbol{\theta}_{0})} \leq J_{ij}(\boldsymbol{\theta}_{0}) + q \frac{\operatorname{tr}\{\mathbf{Q}_{ijk}(\tilde{\boldsymbol{\theta}}) + \mathbf{Q}_{ikj}(\tilde{\boldsymbol{\theta}})\}}{\nu_{k}(\boldsymbol{\theta})\nu_{i}(\boldsymbol{\theta})} \left\{ 1 + c' \frac{q}{\sqrt{m}} \right\}^{3} = J_{ij}(\boldsymbol{\theta}_{0}) + O\left(\frac{q}{\sqrt{m}}\right)$$

Altogether,

$$-\frac{1}{\nu_i(\boldsymbol{\theta}_0)\nu_j(\boldsymbol{\theta}_0)}\frac{\partial^2 \ell_R}{\partial \theta_i \partial \theta_j}(\boldsymbol{\theta}) = J_{ij}(\boldsymbol{\theta}_0) + O_P\left(\frac{q}{\sqrt{m}}\right).$$

The second part of the proof mimics the reasoning of [41]. Let $\mathbf{J}(\boldsymbol{\theta}_0) = \{J_{ij}(\boldsymbol{\theta}_0)\}_{ij}$ and $\mathbf{s}(\boldsymbol{\theta}_0) = \{s_1(\boldsymbol{\theta}_0), s_2(\boldsymbol{\theta}_0), \dots, s_r(\boldsymbol{\theta}_0)\}^\top$ with $s_i(\boldsymbol{\theta}_0) = v_i(\boldsymbol{\theta}_0)^{-1} \partial \ell_R / \partial \theta_i|_{\boldsymbol{\theta}_0}$ and define $\boldsymbol{\theta}'$ such that

$$\nu(\boldsymbol{\theta}_0) \circ (\boldsymbol{\theta}' - \boldsymbol{\theta}_0) = \mathbf{J}(\boldsymbol{\theta}_0)^{-1} \mathbf{s}(\boldsymbol{\theta}_0).$$

We will show that there exists a $\hat{\theta}$ so that

$$|v_i(\boldsymbol{\theta}_0)|\theta_i - \theta_{0,i}| \leq |v_i(\boldsymbol{\theta}_0)|\theta_i - \theta_i'| + |v_i(\boldsymbol{\theta}_0)|\theta_i' - \theta_{0,i}|$$

By (ii) and (iii), the last term is $v_i(\theta_0)|\theta'_i - \theta_{0,i}| = O_P(1)$. It remains to show that $v_i(\theta_0)|\hat{\theta}_i - \theta'_i| = O_P(1)$. This is shown by proving that a maximum of (5) is attained in a region close to θ' .

Now, for any $\theta \in \Theta_q$,

$$\ell_{R}(\boldsymbol{\theta}) = \ell_{R}(\boldsymbol{\theta}_{0}) + \sum_{i=1}^{r} \nu_{i}(\boldsymbol{\theta}_{0})(\theta_{i} - \theta_{0,i})s_{i}(\boldsymbol{\theta}_{0}) - \frac{1}{2}\sum_{i=1}^{r}\sum_{j=1}^{r} \nu_{i}(\boldsymbol{\theta}_{0})(\theta_{i} - \theta_{0,i})\nu_{j}(\boldsymbol{\theta}_{0})(\theta_{j} - \theta_{0,j})J_{ij}(\boldsymbol{\theta}_{0}) + R(\boldsymbol{\theta}, \tilde{\boldsymbol{\theta}})$$

$$= \ell_{R}(\boldsymbol{\theta}_{0}) + \frac{1}{2}\mathbf{s}(\boldsymbol{\theta}_{0})^{\top}\mathbf{J}(\boldsymbol{\theta}_{0})^{-1}\mathbf{s}(\boldsymbol{\theta}_{0}) - \frac{1}{2}\sum_{i=1}^{r}\sum_{j=1}^{r}\nu_{i}(\boldsymbol{\theta}_{0})(\theta_{i}' - \theta_{i})\nu_{j}(\boldsymbol{\theta}_{0})(\theta_{j}' - \theta_{j})J_{ij}(\boldsymbol{\theta}_{0}) + R(\boldsymbol{\theta}, \tilde{\boldsymbol{\theta}}),$$
(19)

where $R(\boldsymbol{\theta}, \tilde{\boldsymbol{\theta}}) = -\frac{1}{2} \sum_{i=1}^{r} \sum_{j=1}^{r} v_i(\boldsymbol{\theta}_0)(\theta_i - \theta_{0,i})v_j(\boldsymbol{\theta}_0)(\theta_j - \theta_{0,j})\zeta(\tilde{\boldsymbol{\theta}}, \boldsymbol{\theta}_0)$ for some $\tilde{\theta}_i = \theta_{0,i} + t(\theta_i - \theta_{0,i})$ where $t \in [0, 1]$. Now consider the set $\Theta'_{\epsilon_n} = \{\boldsymbol{\theta} : v_i(\boldsymbol{\theta}_0)|\theta_i - \theta'_i| < \epsilon_n\}$ and its boundary $\partial \Theta'_{\epsilon_n}$ and $\overline{\Theta}'_{\epsilon_n} = \Theta'_{\epsilon_n} \cup \partial \Theta'_{\epsilon_n}$. For any $\boldsymbol{\theta} \in \overline{\Theta}'_{\epsilon_n}$ with $\epsilon_n \to 0$,

$$\nu_i(\boldsymbol{\theta}_0)|\theta_i - \theta_{0,i}| \le \nu_i(\boldsymbol{\theta}_0)|\theta_i - \theta_i'| + \nu_i(\boldsymbol{\theta}_0)|\theta_i' - \theta_{0,i}| = O_P(1)$$

that is for all $\varepsilon > 0$ there exists a constant *C* and N_{ε} such that for all $n \ge N_{\varepsilon}$: $1 - \varepsilon \le \inf_{\theta_0 \in \Theta} \Pr_{\theta_0} \{ v_i(\theta_0) | \theta_i - \theta'_i | < C \}$. Since $q \to \infty$, there exists $N \ge N_{\varepsilon}$ such that q > C. Together, for any $\theta \in \overline{\Theta}'_{\epsilon_n}$ and any $\varepsilon > 0$ there exists $N \in \mathbb{N}$, such that for all $n \ge N$:

$$1 - \varepsilon \leq \inf_{\boldsymbol{\theta}_0 \in \Theta} \Pr_{\boldsymbol{\theta}_0} \left\{ \nu_i(\boldsymbol{\theta}_0) | \theta_i - \theta_i' | < C \right\} \leq \inf_{\boldsymbol{\theta}_0 \in \Theta} \Pr_{\boldsymbol{\theta}_0} \left\{ \nu_i(\boldsymbol{\theta}_0) | \theta_i - \theta_i' | < q \right\},$$

i.e., $\inf_{\theta_0 \in \Theta} \Pr_{\theta_0}(\overline{\Theta}'_{\epsilon_n} \subset \Theta_q) \to 1$. This implies that

$$\begin{split} \inf_{\boldsymbol{\theta}_{0}\in\Theta} \operatorname{Pr}_{\boldsymbol{\theta}_{0}} \left\{ \sup_{\boldsymbol{\theta}\in\partial\Theta_{\epsilon_{n}}} |R(\boldsymbol{\theta},\tilde{\boldsymbol{\theta}})| \leq \sup_{\boldsymbol{\theta}\in\Theta_{q}} |R(\boldsymbol{\theta},\tilde{\boldsymbol{\theta}})| \right\} \\ &\geq \inf_{\boldsymbol{\theta}_{0}\in\Theta} \operatorname{Pr}_{\boldsymbol{\theta}_{0}} \left\{ \sup_{\boldsymbol{\theta}\in\partial\Theta_{\epsilon_{n}}} |R(\boldsymbol{\theta},\tilde{\boldsymbol{\theta}})| \leq \sup_{\boldsymbol{\theta}\in\Theta_{q}} |R(\boldsymbol{\theta},\tilde{\boldsymbol{\theta}})| \left| \left| \partial\Theta_{\epsilon_{n}}' \subset \Theta_{q} \right\} \inf_{\boldsymbol{\theta}_{0}\in\Theta} \operatorname{Pr}_{\boldsymbol{\theta}_{0}} \left(\partial\Theta_{\epsilon_{n}}' \subset \Theta_{q} \right) \right. \\ &+ \inf_{\boldsymbol{\theta}_{0}\in\Theta} \operatorname{Pr}_{\boldsymbol{\theta}_{0}} \left\{ \sup_{\boldsymbol{\theta}\in\partial\Theta_{\epsilon_{n}}} |R(\boldsymbol{\theta},\tilde{\boldsymbol{\theta}})| \leq \sup_{\boldsymbol{\theta}\in\Theta_{q}} |R(\boldsymbol{\theta},\tilde{\boldsymbol{\theta}})| \left| \left| \partial\Theta_{\epsilon_{n}}' \not\subset \Theta_{q} \right\} \inf_{\boldsymbol{\theta}_{0}\in\Theta} \operatorname{Pr}_{\boldsymbol{\theta}_{0}} \left(\partial\Theta_{\epsilon_{n}}' \not\subset \Theta_{q} \right) \rightarrow 1 \end{split}$$

Second-to-last, consider

$$\delta(\epsilon_n) = \min_{\boldsymbol{\theta} \in \partial \Theta'_{\epsilon_n}} \frac{1}{2} \sum_{i=1}^r \sum_{j=1}^r \nu_i(\boldsymbol{\theta}_0)(\theta'_i - \theta_i)\nu_j(\boldsymbol{\theta}_0)(\theta'_j - \theta_j)J_{ij}(\boldsymbol{\theta}_0) = \frac{\epsilon_n^2}{2} \mathbf{1}_r^\top \mathbf{J}(\boldsymbol{\theta}_0)\mathbf{1}_r \ge c'\frac{r\epsilon_n^2}{2} > 0$$

by (iii). Note that $\delta(\epsilon_n)$ is not stochastic, increasing in ϵ_n and $\delta(0) = 0$. Further, by (iv), $2 \sup_{\theta \in \Theta_q} |R(\theta, \tilde{\theta})| = O_P(q^3/\sqrt{m})$, i.e., for all $\varepsilon > 0$ there exists a constant C' and N'_{ε} such that for all $n \ge N'_{\varepsilon}$ it holds that $1 - \varepsilon \le \inf_{\theta_0 \in \Theta} \Pr_{\theta_0} \{2 \sup_{\theta \in \Theta_q} |R(\theta, \tilde{\theta})| < C'q^3/\sqrt{m}\}$. Now set $\epsilon_n \to 0$ so that it exists $N' > N'_{\varepsilon}$ such that for all $n > N' : \epsilon_n^2 rc'/2 > C'q^3/\sqrt{m}$. This gives that for all n > N':

$$\begin{split} 1 - \varepsilon &\leq \inf_{\theta_0 \in \Theta} \Pr_{\theta_0} \left\{ 2 \sup_{\theta \in \Theta_q} |R(\theta, \tilde{\theta})| < C' \frac{q^3}{\sqrt{m}} \right\} \leq \inf_{\theta_0 \in \Theta} \Pr_{\theta_0} \left\{ 2 \sup_{\theta \in \Theta_q} |R(\theta, \tilde{\theta})| < c' \frac{r\epsilon_n^2}{2} \right\} \\ &\leq \inf_{\theta_0 \in \Theta} \Pr_{\theta_0} \left\{ 2 \sup_{\theta \in \Theta_q} |R(\theta, \tilde{\theta})| < \delta(\epsilon_n) \right\}. \end{split}$$

Finally, this implies that for $\theta' \in \Theta_a$ and by (19)

$$\begin{split} \lim_{n \to \infty} \inf_{\theta_0 \in \Theta} \Pr_{\theta_0} \left\{ \ell_R(\theta') > \max_{\theta \in \partial \Theta'_{\epsilon_n}} \ell_R(\theta) \right\} &= \lim_{n \to \infty} \inf_{\theta_0 \in \Theta} \Pr_{\theta_0} \left\{ \delta(\epsilon_n) > \max_{\theta \in \partial \Theta'_{\epsilon_n}} R(\theta, \tilde{\theta}) - R(\theta', \tilde{\theta}') \\ &\geq \lim_{n \to \infty} \inf_{\theta_0 \in \Theta} \Pr_{\theta_0} \left\{ \delta(\epsilon_n) > 2 \sup_{\theta \in \Theta_q} |R(\theta, \tilde{\theta})| \right\} = 1, \end{split}$$

i.e., with probability going to one, ℓ_R attains a local maximum in Θ'_{ϵ_n} . Denote $\hat{\theta}$ as a local maximum of ℓ_R in Θ'_{ϵ_n} if one exists and set $\hat{\theta} \notin \Theta'_{\epsilon_n}$ otherwise. Then, above statement gives that for all $\varepsilon > 0$ there exists N''_{ε} such that for all $n \ge N''_{\varepsilon}$:

 $\inf_{\theta_0 \in \Theta} \Pr_{\theta_0}(\hat{\theta} \in \Theta'_{\epsilon_n}) \ge 1 - \varepsilon$. Now, for any constant C'' > 0 and any $\varepsilon > 0$ let N'' be such that $C'' > \epsilon_{N''}$ and $N'' > N''_{\varepsilon}$. Then, for all n > N'',

$$\begin{split} \sup_{\boldsymbol{\theta}_{0}\in\Theta} \Pr_{\boldsymbol{\theta}_{0}} \left\{ \nu_{i}(\boldsymbol{\theta}_{0}) | \hat{\theta}_{i} - \theta_{i}' | \geq C'' \right\} &\leq \sup_{\boldsymbol{\theta}_{0}\in\Theta} \Pr_{\boldsymbol{\theta}_{0}} \left\{ \nu_{i}(\boldsymbol{\theta}_{0}) | \hat{\theta}_{i} - \theta_{i}' | \geq C'', \hat{\boldsymbol{\theta}} \in \Theta_{\epsilon_{n}}' \right\} \\ &+ \sup_{\boldsymbol{\theta}_{0}\in\Theta} \Pr_{\boldsymbol{\theta}_{0}} \left\{ \nu_{i}(\boldsymbol{\theta}_{0}) | \hat{\theta}_{i} - \theta_{i}' | \geq C'', \hat{\boldsymbol{\theta}} \notin \Theta_{\epsilon_{n}}' \right\} \\ &\leq \sup_{\boldsymbol{\theta}_{0}\in\Theta} \Pr_{\boldsymbol{\theta}_{0}} \left\{ \hat{\boldsymbol{\theta}} \notin \Theta_{\epsilon_{n}}' \right\} \leq \epsilon, \end{split}$$

that is $v_i(\boldsymbol{\theta}_0)|\hat{\theta}_{0,i} - \theta'_i| = O_P(1).$

To address the infimum over Θ , we use the following result.

Lemma 2. For $n \in \mathbb{N}$, let (X_n) and (Y_n) be sequences of real-valued continuous random variables with finite variances, in x and y uniformly bounded conditional density $p_{X_n|Y_n=y}(x) = O(1)$ and $X_n = O_P(a_n)$ and $Y_n = O_P(a_nb_n)$ with sequences $(a_n), (b_n) \in \mathbb{R}$ with $b_n = o(1)$. Then, as $n \to \infty$,

$$\Pr(X_n + Y_n \le a_n) = \Pr(X_n \le a_n) + O(b_n).$$

The asymptotic result is clear as convergence in probability implies convergence in distribution. The above result further specifies the rate of convergence.

Proof. First, let $\phi(s, t) = P(X_n + s \le a_n | Y_n = t)$ and consider a Taylor expansion for $f(x) = P(X_n/a_n + x \le 1 | Y_n = t)$ around x = 0, which gives $\phi(s, t) = \phi(0, t) + O(s/a_n)$. This implies $\int_{a_n-t}^{a_n} p_{X_n|Y_n=t}(z) dz = \phi(0, t) - \phi(t, t) = O(t/a_n)$. Using convolution the formula we obtain

$$\Pr(X_{n} + Y_{n} \le a_{n}) = \int_{-\infty}^{a_{n}} p_{X_{n}+Y_{n}}(z) dz = \int_{-\infty}^{a_{n}} \int_{-\infty}^{\infty} p_{X_{n},Y_{n}}(z-t,t) dt dz = \int_{-\infty}^{a_{n}} \int_{-\infty}^{\infty} p_{X_{n}|Y_{n}=t}(z-t) p_{Y_{n}}(t) dt dz$$

$$= \int_{-\infty}^{\infty} p_{Y_{n}}(t) \int_{-\infty}^{a_{n}-t} p_{X_{n}|Y_{n}=t}(z) dz dt = \int_{-\infty}^{\infty} p_{Y_{n}}(t) \left\{ \int_{-\infty}^{a_{n}} p_{X_{n}|Y_{n}=t}(z) dz + \int_{a_{n}-t}^{a_{n}} p_{X_{n}|Y_{n}=t}(z) dz \right\} dt$$

$$= \int_{-\infty}^{\infty} p_{Y_{n}}(t) \left\{ \int_{-\infty}^{a_{n}} p_{X_{n}|Y_{n}=t}(z) dz + O\left(\frac{t}{a_{n}}\right) \right\} dt = \int_{-\infty}^{\infty} \int_{-\infty}^{a_{n}} p_{X_{n},Y_{n}}(z,t) dz dt + O\left\{ E\left(\frac{Y_{n}}{a_{n}}\right) \right\}$$

$$= \int_{-\infty}^{\infty} \int_{-\infty}^{a_{n}} p_{X_{n}}(z) p_{Y_{n}|X_{n}=z}(t) dz dt + O(b_{n}) = \Pr(X_{n} \le a_{n}) + O(b_{n}),$$
(20)

which gives the claim. \Box

The next two results are helpful to prove Theorem 2.

Lemma 3. Let model (2) and (A)-(E) hold and set $m = \min\{rk(\mathbf{H}_1), \ldots, rk(\mathbf{H}_r)\} \rightarrow \infty$ and $\hat{t} = t + \omega > 0$ with

$$t = O\left(p + \left\|\mathbf{C}^{-1/2}\boldsymbol{\Lambda}\mathbf{d}\right\|^{2}\right), \qquad \omega = O_{P}\left(\frac{1}{\sqrt{m}}\left\|\mathbf{C}^{-1/2}\boldsymbol{\Lambda}\mathbf{d}\right\|^{2}\right)$$

Then, there exists a sequence of local maximizers $\hat{\theta}$ of (5) such that

$$\inf_{\substack{\mathbf{d}\in[-1,1]^{p}\\\boldsymbol{\theta}_{0}\in\Theta}}\Pr_{\boldsymbol{\theta}_{0}}\left\{\hat{\mathbf{u}}_{\mathbf{d}}\in E\left(\hat{\mathbf{C}},\hat{t}\right)\right\}=\inf_{\substack{\mathbf{d}\in[-1,1]^{p}\\\boldsymbol{\theta}_{0}\in\Theta}}\Pr_{\boldsymbol{\theta}_{0}}\left\{\mathbf{u}_{\mathbf{d}}\in E\left(\mathbf{C},t\right)\right\}+O\left(\frac{1}{\sqrt{m}}\right)$$

uniformly over $\theta_0 \in \Theta$.

Proof. First, let $\xi = \|\mathbf{C}^{-1/2} \boldsymbol{\Lambda} \mathbf{d}\|^2$ and observe that $\|\mathbf{C}^{1/2} \mathbf{u}_{\mathbf{d}}\|^2 \sim \chi_p^2(\xi)$. This implies that $\|\mathbf{C}^{1/2} \mathbf{u}_{\mathbf{d}}\|^2 = O_P(1+\xi)$ for all $\boldsymbol{\theta}_0$. Now, consider its derivative with respect to θ_i , $i \in \{1, ..., r\}$.

$$\frac{\partial}{\partial \theta_i} \| \mathbf{C}^{1/2} \mathbf{u_d} \|^2 = \| \boldsymbol{\Omega}_i^{1/2} (\mathbf{w} - \boldsymbol{\Lambda} \mathbf{d}) \|^2 + 2(\mathbf{w} - \boldsymbol{\Lambda} \mathbf{d})^\top \mathbf{C}^{-1} \frac{\partial \mathbf{w}}{\partial \theta_i}$$

where $\boldsymbol{\Omega}_i = \mathbf{C}^{-1} \mathbf{X}^{\mathsf{T}} \mathbf{V}^{-1} \mathbf{H}_i \mathbf{V}^{-1} \mathbf{X} \mathbf{C}^{-1} / n$ as $\partial \mathbf{C}^{-1} / \partial \theta_i = \boldsymbol{\Omega}_i$. Now,

$$\mathsf{E}\left(\frac{\partial}{\partial\theta_{i}}\|\mathbf{C}^{1/2}\mathbf{u}_{\mathbf{d}}\|^{2}\right) = \|\boldsymbol{\Omega}_{i}^{1/2}\boldsymbol{\Lambda}\mathbf{d}\|^{2} - \mathsf{tr}(\mathbf{C}\boldsymbol{\Omega}_{i}) \leq \|\boldsymbol{\Omega}_{i}^{1/2}\boldsymbol{\Lambda}\mathbf{d}\|^{2} + \mathsf{tr}(\mathbf{C}\boldsymbol{\Omega}_{i}).$$

Let $\mathbf{A}_i = \mathbf{V}^{-1/2} \mathbf{H}_i \mathbf{V}^{-1/2}$ and $\mathbf{B} = \mathbf{V}^{-1/2} \mathbf{X} \mathbf{C}^{-1} (\mathbf{\Lambda} \mathbf{d} \mathbf{d}^\top \mathbf{\Lambda} + \mathbf{C}) \mathbf{C}^{-1} \mathbf{X}^\top \mathbf{V}^{-1/2} / n$. Both \mathbf{A}_i and \mathbf{B} are positive semi-definite. Since $rk(\mathbf{B}) = p \text{ and } \eta_1(\mathbf{B}) = O(1 + \xi),$

$$\mathsf{E}\left(\frac{\partial}{\partial\theta_{i}}\|\mathbf{C}^{1/2}\mathbf{u}_{\mathbf{d}}\|^{2}\right) \leq \operatorname{tr}(\mathbf{A}_{i}\mathbf{B}) \leq \sum_{j=1}^{p}\eta_{j}(\mathbf{A}_{i})\eta_{j}(\mathbf{B}) = O\left\{(1+\xi)\sum_{j=1}^{p}\eta_{j}(\mathbf{A}_{i})\right\}$$

for all θ_0 . Similarly, lengthy calculations give that

$$\operatorname{Var}\left(\frac{\partial}{\partial\theta_{i}}\|\mathbf{C}^{1/2}\mathbf{u}_{\mathbf{d}}\|^{2}\right) = 2\operatorname{tr}(\boldsymbol{\Omega}_{i}\mathbf{C}\boldsymbol{\Omega}_{i}\mathbf{C}) - 4\|\mathbf{C}^{1/2}\boldsymbol{\Omega}_{i}\boldsymbol{\Lambda}\mathbf{d}\|^{2} + \frac{4}{n}\|\mathbf{V}^{-1/2}\mathbf{H}_{i}\mathbf{V}^{-1}\mathbf{X}\mathbf{C}^{-1}\boldsymbol{\Lambda}\mathbf{d}\|^{2}$$
$$\leq 2\operatorname{tr}(\boldsymbol{\Omega}_{i}\mathbf{C}\boldsymbol{\Omega}_{i}\mathbf{C}) + 4\|\mathbf{C}^{1/2}\boldsymbol{\Omega}_{i}\boldsymbol{\Lambda}\mathbf{d}\|^{2} + \frac{4}{n}\|\mathbf{V}^{-1/2}\mathbf{H}_{i}\mathbf{V}^{-1}\mathbf{X}\mathbf{C}^{-1}\boldsymbol{\Lambda}\mathbf{d}\|^{2}.$$

Proceeding as above, for all θ_0 ,

$$\operatorname{Var}\left(\frac{\partial}{\partial\theta_{i}}\|\mathbf{C}^{1/2}\mathbf{u}_{\mathbf{d}}\|^{2}\right) = O\left[(1+\xi)\left\{\sum_{j=1}^{p}\eta_{j}(\mathbf{A}_{i})\right\}^{2}\right].$$

Noting that $c\eta_i\{\mathbf{A}_i(\mathbf{1}_r)\} \ge \theta_{0,i}\eta_i\{\mathbf{A}_i(\boldsymbol{\theta}_0)\} \ge c^{-1}\eta_i\{\mathbf{A}_i(\mathbf{1}_r)\}, j \in \{1, \dots, n\}$, Chebyshev's inequality gives the representation

$$\frac{\partial}{\partial \theta_i} \left\| \mathbf{C}^{1/2} \mathbf{u_d} \right\|^2 = O_P \left\{ \frac{\| \mathbf{C}^{1/2} \mathbf{u_d} \|^2}{\theta_{0,i}} \right\} = O_P \left\{ \| \mathbf{C}^{1/2} \mathbf{u_d} \|^2 \frac{\nu_i}{\sqrt{m}} \right\},\tag{21}$$

for all θ_0 . By Theorem 1, a Taylor expansion for $\|\hat{\mathbf{C}}^{1/2}\hat{\mathbf{u}}_{\mathbf{d}}\|^2$ around θ_0 gives

$$\|\hat{\mathbf{C}}^{1/2}\hat{\mathbf{u}}_{\mathbf{d}}\|^{2} = \|\mathbf{C}^{1/2}\mathbf{u}_{\mathbf{d}}\|^{2} + O_{P}\left\{ (\hat{\boldsymbol{\theta}} - \boldsymbol{\theta}_{0})^{\top} \frac{\partial}{\partial \boldsymbol{\theta}} \|\mathbf{C}^{1/2}\mathbf{u}_{\mathbf{d}}\|^{2} \right\} = \|\mathbf{C}^{1/2}\mathbf{u}_{\mathbf{d}}\|^{2} \left\{ 1 + O_{P}(m^{-1/2}) \right\},$$

which holds for all θ_0 . Eventually, let $X_n = \|\mathbf{C}^{1/2}\mathbf{u}_d\|^2/(1+\xi)$, $Y_n = O_P\{m^{-1/2}X_n + m^{-1/2}\xi/(1+\xi)\} = O_P\{b_nX_n\}$ for all θ_0 with $b_n = m^{-1/2}$ and $a_n = t(1 + \xi)^{-1}$. Then,

$$\inf_{\boldsymbol{\theta}_{0}\in\Theta}\Pr_{\boldsymbol{\theta}_{0}}\left(\left\|\hat{\mathbf{C}}^{1/2}\hat{\mathbf{u}}_{\mathbf{d}}\right\|^{2}\leq\hat{t}\right)=\inf_{\boldsymbol{\theta}_{0}\in\Theta}\Pr_{\boldsymbol{\theta}_{0}}\left\{X_{n}+Y_{n}\leq a_{n}\right\}$$

Finally, Lemma 2 gives the claim. \Box

Lemma 4. Let model (2) and (A)–(B) hold. Then, for t > 0,

$$\operatorname{argmin}_{\mathbf{d}\in\{-1,1\}^p} \operatorname{Pr}_{\boldsymbol{\theta}_0} \left\{ \mathbf{u}_{\mathbf{d}} \in E(\mathbf{C}, t) \right\} = \operatorname{argmax}_{\mathbf{d}\in\{-1,1\}^p} \left\| \mathbf{C}^{-1/2} \boldsymbol{\Lambda} \mathbf{d} \right\|^2.$$

This result is given in [9, Prop. 4].

Proof of Theorem 2. Consider $\tau = \max_{\mathbf{d}} \chi_{p,1-\alpha}^2(\xi)$ with $\xi = \|\mathbf{C}^{-1/2} \Lambda \mathbf{d}\|^2$. Since for $X \sim \chi_p^2(\xi)$ it holds $X = O_P(1+\xi)$ for all θ_0 and by the definition of the quantile $\Pr(X \le \tau) = 1 - \alpha$ it follows that $\tau = O(1+\xi)$ for all θ_0 as well. Now we proceed similarly to the proof of Lemma 3. A Taylor expansion for $\|\hat{\mathbf{C}}^{-1/2} \Lambda \mathbf{d}\|^2$ around θ_0 gives for $\Omega_i = 0$

 $\mathbf{C}^{-1}\mathbf{X}^{\top}\mathbf{V}^{-1}\mathbf{H}_{i}\mathbf{V}^{-1}\mathbf{X}\mathbf{C}^{-1}/n$ that

$$\|\hat{\mathbf{C}}^{-1/2}\boldsymbol{\Lambda}\mathbf{d}\|^{2} = \|\mathbf{C}^{-1/2}\boldsymbol{\Lambda}\mathbf{d}\|^{2} + O_{P}\left\{\sum_{i=1}^{p} (\hat{\theta}_{i} - \theta_{0,i}) \|\boldsymbol{\Omega}_{i}^{1/2}\boldsymbol{\Lambda}\mathbf{d}\|^{2}\right\} = \|\mathbf{C}^{-1/2}\boldsymbol{\Lambda}\mathbf{d}\|^{2} + O_{P}(m^{-1/2}\xi)$$

for all θ_0 and together with the first argument it follows that $\hat{\tau} = \tau + O_P(m^{-1/2}\xi)$ for all θ_0 . By Lemma 3 it is ensured that the coverage is attained uniformly for both β_0 and θ_0 ,

$$\inf_{\substack{\boldsymbol{\beta}_0 \in \mathbb{R}^p\\\boldsymbol{\theta}_0 \in \Theta}} \Pr_{\boldsymbol{\beta}_0, \boldsymbol{\theta}_0} \left\{ \sqrt{n} \left(\hat{\boldsymbol{\beta}}_L - \boldsymbol{\beta}_0 \right) \in E\left(\hat{\mathbf{C}}, \hat{\boldsymbol{\tau}} \right) \right\} = \inf_{\substack{\mathbf{d} \in \{-1, 1\}^p\\\boldsymbol{\theta}_0 \in \Theta}} \Pr_{\boldsymbol{\theta}_0} \left\{ \mathbf{u}_{\mathbf{d}} \in E\left(\mathbf{C}, \boldsymbol{\tau} \right) \right\} + O(m^{-1/2}).$$

By Lemma 4, this minimum is in fact attained for the $\mathbf{d} \in \{-1, 1\}^p$ for which τ ensures nominal coverage, since $\|\mathbf{C}^{1/2}\mathbf{u}_{\mathbf{d}}\|^2 \sim \chi_p^2(\xi)$. This proves the claim. \Box

CRediT authorship contribution statement

Peter Kramlinger: Methodology Writing, Software. Ulrike Schneider: Methodology, Writing. Tatyana Krivobokova: Methodology, Writing.

Acknowledgments

This contribution has been partially created while the authors Peter Kramlinger and Tatyana Krivobokova were at the University of Göttingen. The authors gratefully acknowledge the funding by the German Research Association (DFG) via Research Training Group 1644 "Scaling Problems in Statistics" and thank anonymous referees for their insightful comments and suggestions.

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